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(57) Abstract: The present application discloses an antibody, or fragment thereof, for the diagnosis, treatment or prevention of cancers wherein the antibody specifically binds to the PSMGFR peptide (SEQ ID NO:2) or a fragment thereof of the peptide.

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## ANTI-VARIABLE MUC1\* ANTIBODIES AND USES THEREOF

### BACKGROUND OF THE INVENTION

#### [0001] 1. Field of the Invention:

[0002] The present application relates to human, humanized and non-human anti-MUC1\* antibodies and methods of making and using them. The present application also relates to using an immune cell transfected or transduced with a cleavage enzyme for the treatment of cancer. The present invention also relates to using an immune cell transfected or transduced with a CAR and another protein for the treatment of cancer.

#### [0003] 2. General Background and State of the Art:

[0004] We previously discovered that a cleaved form of the MUC1 (SEQ ID NO:1) transmembrane protein is a growth factor receptor that drives the growth of over 75% of all human cancers. The cleaved form of MUC1, which we called MUC1\* (pronounced muk 1 star), is a powerful growth factor receptor. Cleavage and release of the bulk of the extracellular domain of MUC1 unmask a binding site for activating ligands dimeric NME1, NME6, NME7, NME7<sub>AB</sub>, NME7-X1 or NME8. It is an ideal target for cancer drugs as it is aberrantly expressed on over 75% of all cancers and is likely overexpressed on an even higher percentage of metastatic cancers (Mahanta et al. (2008) A Minimal Fragment of MUC1 Mediates Growth of Cancer Cells. PLoS ONE 3(4): e2054. doi:10.1371/ journal.pone.0002054; Fessler et al. (2009), "MUC1\* is a determinant of trastuzumab (Herceptin) resistance in breast cancer cells," Breast Cancer Res Treat. 118(1):113-124). After MUC1 cleavage most of its extracellular domain is shed from the cell surface. The remaining portion has a truncated extracellular domain that comprises most or all of the primary growth factor receptor sequence called PSMGFR (SEQ ID NO:2).

[0005] Antibodies are increasingly used to treat human diseases. Antibodies generated in non-human species have historically been used as therapeutics in humans, such as horse antibodies. More recently, antibodies are engineered or selected so that they contain mostly, or all, human sequences in order to avoid a generalized rejection of the foreign antibody. The process of engineering recognition fragments of a non-human antibody into a human antibody is generally called 'humanizing'. The amount of non-human sequences that are used to replace the



human antibody sequences determines whether they are called chimeric, humanized or fully human.

**[0006]** Alternative technologies exist that enable generation of humanized or fully human antibodies. These strategies involve screening libraries of human antibodies or antibody fragments and identifying those that bind to the target antigen, rather than immunizing an animal with the antigen. Another approach is to engineer the variable region(s) of an antibody into an antibody-like molecule. Another approach involves immunizing a humanized animal. The present invention is intended to also encompass these approaches for use with recognition fragments of antibodies that the inventors have determined bind to the extracellular domain of MUC1\*.

**[0007]** In addition to treating patients with an antibody, cancer immunotherapies have recently been shown to be effective in the treatment of blood cancers. One cancer immunotherapy, called CAR T (chimeric antigen receptor T cell) therapy, engineers a T cell so that it expresses a chimeric receptor having an extra cellular domain that recognizes a tumor antigen, a transmembrane domain and cytoplasmic tail comprising T cell signaling and co-stimulatory components (Dai H, Wang Y, Lu X, Han W. (2016) Chimeric Antigen Receptors Modified T-Cells for Cancer Therapy. J Natl Cancer Inst. 108(7): djv439). Such receptor is composed of a single chain antibody fragment (scFv) that recognizes a tumor antigen, linked to a T cell transmembrane, signaling domain and co-stimulatory domain or domains. Upon binding of the receptor to a cancer associated antigen, a signal is transmitted resulting in T-cell activation, propagation and the targeted killing of the cancer cells. In practice, T cells are isolated from a patient or donor and transduced with a CAR, expanded and then injected back into the patient. If from a donor, the immune cells may be mutated or engineered such that they do not induce graft versus host disease in the recipient. When the CAR T cells bind to the antigen on a cancer cell, the CAR T cells attack the cancer cells and then expand that population of T cells.

**[0008]** Thus far, CAR T therapies have been very successful in the treatment of blood cancers but as yet have not shown efficacy against solid tumors in humans. Because most blood cancers are B cell malignancies, the CAR T cells can just eliminate all of the patient's B cells without causing serious harm to the patient. There is no B cell equivalent in solid tumors. Most tumor associated antigens are also expressed on normal tissues; they are just expressed at a higher level in cancerous tissues. Thus, the challenge is to develop an antibody that recognizes

an epitope on a tumor associated antigen that is somehow different in the context of the tumor compared to normal tissue. To further minimize the risk of off-tumor/on-target killing of normal tissues, the antibody should recognize and bind to cancerous tissues at least two-times more than normal tissues. Antibodies that are not so cancer selective may be used therapeutically if they are inducibly expressed at the tumor site.

**[0009]** Another cancer therapy that incorporates cancer selective antibodies is Bi-specific T cell Engagers, also called BiTEs. The BiTE approach attempts to eliminate the CAR T associated risk of off-tumor/on-target effects. Unlike CAR T, BiTEs are bispecific antibodies that should not pose any greater risk than regular antibody-based therapies. However, unlike typical anti-cancer antibodies that bind to and block a cancer antigen, BiTEs are designed to bind to an antigen on the tumor cell and simultaneously bind to an antigen on an immune cell, such as a T cell. In this way, a BiTE recruits the T cell to the tumor. BiTEs are engineered proteins that simultaneously bind to a cancer associated antigen and a T-cell surface protein such as CD3-epsilon. BiTEs are antibodies made by genetically linking the scFv's of an antibody that binds to a T cell antigen, like anti-CD3-epsilon to a scFv of a therapeutic monoclonal antibody that binds to a cancer antigen (Patrick A. Baeuerle, and Carsten Reinhardt (2009) Bispecific T-cell engaging antibodies for cancer therapy. Cancer Res. 69(12):4941-4944). A drawback of BiTE technology is that, unlike CAR T cells, they do not expand in the patient, so have limited persistence.

**[0010]** Yet another cancer therapy that incorporates cancer selective antibodies is antibody drug conjugate, also called ADC, technology. In this case, a toxin, or a precursor to a toxin, is linked to a cancer selective antibody. Unlike CAR T cells that use the CD8 positive T cell's natural killing to kill cancer cells, ADCs carry a toxic payload to the tumor. Drawbacks of ADCs include the potential of delivering the toxic payload to normal cells and that most ADCs require binding to a cell surface molecule which then gets internalized after binding, with an approximate 10,000 surface molecule required for resultant cell death.

## **SUMMARY OF THE INVENTION**

**[0011]** In one aspect, the present invention is directed to a non-human, human or humanized anti-MUC1\* antibody or antibody fragment or antibody-like protein that binds to a region on extracellular domain of MUC1 isoform or cleavage product that is devoid of the tandem repeat

domains. The non-human, human or humanized anti-MUC1\* antibody or antibody fragment or antibody-like protein may specifically bind to

**[0012]** (i) PSMGFR region of MUC1;

**[0013]** (ii) PSMGFR peptide;

**[0014]** (iii) a peptide having amino acid sequence of QFNQYKTEAASRYNLTISDVSVDVPFPFSAQSGA (N-10) (SEQ ID NO:3)

**[0015]** (iv) a peptide having amino acid sequence of

**[0016]** ASRYNLTISDVSVDVPFPFSAQSGA (N-19) (SEQ ID NO:4)

**[0017]** (v) a peptide having amino acid sequence of

**[0018]** NLTISDVSVDVPFPFSAQSGA (N-23) (SEQ ID NO:5)

**[0019]** (vi) a peptide having amino acid sequence of

**[0020]** ISDVSVDVPFPFSAQSGA (N-26) (SEQ ID NO:6)

**[0021]** (vii) a peptide having amino acid sequence of

**[0022]** SVSDVPFPFSAQSGA (N-30) (SEQ ID NO:7)

**[0023]** (viii) a peptide having amino acid sequence of

**[0024]** QFNQYKTEAASRYNLTISDVSVDVPFPFS (N-10/C-5) (SEQ ID NO:8)

**[0025]** (ix) a peptide having amino acid sequence of

**[0026]** ASRYNLTISDVSVDVPFPFS (N-19/C-5) (SEQ ID NO:9)

**[0027]** (x) a peptide having amino acid sequence of

**[0028]** FPFSAQSGA (SEQ ID NO:10)

**[0029]** The non-human, human or humanized antibody may be IgG1, IgG2, IgG3, IgG4 or IgM. The human or humanized antibody fragment or antibody-like protein may be scFv or scFv-Fc.

**[0030]** The murine, camelid, human or humanized antibody, antibody fragment or antibody-like protein as in above may comprise a heavy chain variable region and light chain variable region which is derived from mouse monoclonal MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, and H11 antibody, and has at least 80%, 90% or 95% or 98% sequence identity to the mouse monoclonal MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, and H11 antibody. The heavy chain variable region of CDR1 and CDR2 may have at least 90% or 95% or 98% sequence identity to the particularly indicated antibody heavy chain variable region sequence set forth in the present

application in the sequence listing, and the light chain variable region of CDR1 and CDR2 may have at least 90% or 95% or 98% sequence identity to the particularly indicated antibody heavy chain variable region sequence set forth in the present application in the sequence listing section. The heavy chain variable region of CDR3 may have at least 80% or 85% or 90% sequence identity to the particularly indicated antibody heavy chain variable region sequence set forth in the present application in the sequence listing, and the light chain variable region of CDR3 may have at least 80% or 85% or 90% sequence identity to the particularly indicated antibody heavy chain variable region sequence set forth in the present application in the sequence listing section.

**[0031]** The murine, camelid, human or humanized antibody, antibody fragment or antibody-like protein according to above may include complementarity determining regions (CDRs) in the heavy chain variable region and light chain variable region having at least 90% or 95% or 98% sequence identity to the particularly indicated antibody heavy chain CDR1, CDR2 or CDR3 region and light chain CDR1, CDR2 or CDR3 region sequences set forth in the present application in the sequence listing section.

**[0032]** In another aspect, the present invention is directed to an anti-MUC1\* extracellular domain antibody or anti-N-10 antibody, which may be any of the antibodies described above, comprised of sequences represented by humanized IgG2 heavy chain, or humanized IgG1 heavy chain, paired with humanized Kappa light chain, or humanized Lambda light chain. The humanized IgG2 heavy chain may be SEQ ID NOS:53, humanized IgG1 heavy chain may be SEQ ID NO:57, humanized Kappa light chain may be SEQ ID NO:108, and humanized Lambda light chain may be SEQ ID NO:112, or a sequence having 90%, 95% or 98% sequence identity thereof.

**[0033]** In another aspect, the invention is directed to an anti-MUC1\* extracellular domain antibody or anti-N-10 antibody comprised of sequences of a humanized MN-C2 represented by humanized IgG1 heavy chain, humanized IgG2 heavy chain, paired with humanized Lambda light chain, and humanized Kappa light chain.

**[0034]** In another aspect, the invention is directed to an anti-MUC1\* extracellular domain antibody or anti-N-10 antibody comprised of sequences of a humanized MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11 represented by humanized IgG1 heavy chain or humanized IgG2 heavy chain, paired with humanized Lambda light chain, or humanized Kappa light chain.

**[0035]** In another aspect, the invention is directed to an antibody that is “like” MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11 in that they have the same or very similar pattern of binding to subsets of peptides derived from the PSMGFR peptide, also do not recognize a linear epitope, competitively inhibit the binding of NME1 or NME7<sub>AB</sub> to MUC1\*, recognize a MUC1 transmembrane cleavage product produced by cleavage by MMP9 or contain CDR sequences that are at least 80% homologous to the MNE6, MN-C2, MN-18G12, MN-20A10, MN-25E6, MN-28F9, MN-5C6F3, MN-3C2B1, and MN-1E4 CDR consensus sequences.

**[0036]** In another aspect, the invention is directed to an antibody that binds to the extracellular domain of a MUC1 that is devoid of the tandem repeat domain, which may be a cleavage product. In one aspect of the invention, the antibody binds to a peptide having the sequence of QFNQYKTEAASRYNLTISDVSVSDVPFPFSAQSGA (N-10). In one aspect of the invention, the antibody binds to a peptide having the sequence of ASRYNLTISDVSVSDVPFPFSAQSGA (N-19). In one aspect of the invention, the antibody binds to a peptide having the sequence of SVSDVPFPFSAQSGA (N-30). In one aspect of the invention, the antibody binds to a peptide having the sequence of FPFSAQSGA (N-36). Examples of such antibodies include but are not limited to monoclonal antibodies MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, and H11. The heavy chain and light chain complementary determining region sequences for these antibodies are set forth in the present application in the sequence listing section.

**[0037]** In one aspect of the invention, one or more of these antibodies is administered to a patient diagnosed with or at risk of developing a cancer. The antibody may be human or humanized. The antibody may be murine or camelid. The antibody may be bivalent or monovalent. The antibody may be a fragment, including a single chain fragment, scFv, of one of the antibodies. The antibody or antibody fragment may be administered directly to the patient or incorporated into a bispecific antibody, a bispecific T cell engager, BiTE, or an antibody drug conjugate, ADC. The antibody or antibody fragment may be incorporated into a T cell receptor, TCR. The sequence of the antibody or antibody fragment may be incorporated into a chimeric antigen receptor, a “CAR”, or other similar entity, then introduced into an immune cell, *ex vivo*, then administered to a patient diagnosed with or at risk of developing a cancer. The immune cell, which may be a T cell or natural killer cell, may be derived from a donor or from the patient. In

one aspect the immune cell is derived from a stem cell that has been directed to differentiate to that immune cell type in vitro. In one aspect, the antibody or a CAR containing sequences of the antibody may be expressed off of an inducible promoter. In one case the antibody or the CAR is expressed upon activation of the T cell or other immune cell. In one instance, the antibody or the CAR of the invention is expressed off of an NFAT response element. In another instance, CAR recognition of a target tumor cell activates the immune cell, leading to NFAT inducible expression of a cytokine, such as IL-12 or IL-18, or expression of a checkpoint inhibitor such as a PD1 inhibitor or a PDL-1 inhibitor. In yet another aspect, CAR recognition of a target tumor cell activates the immune cell, leading to NFAT inducible expression of a second CAR that contains sequences of a second antibody.

**[0038]** In another aspect, the invention is directed to a murine, camelid, human, humanized anti-MUC1\* antibody or antibody fragment or antibody-like protein that binds to the N-10 peptide, according to above, which inhibits the binding of NME protein to MUC1\*. The NME may be NME1, NME6, NME7<sub>AB</sub>, NME7-X1, NME7 or NME8.

**[0039]** In yet another aspect, the invention is directed to a single chain variable fragment (scFv) comprising a heavy and light chain variable regions connected via a linker, further comprising CDRs of antibodies that bind to MUC1\* extracellular domain. The CDRs may be derived from MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, and H11. The scFv may be one that possesses the SEQ ID NOS:233, 235 and 237 (MN-E6); SEQ ID NOS:239, 241, and 243 (MN-C2)

**[0040]** In still another aspect, the invention is directed to a chimeric antigen receptor (CAR) comprising a scFv or a humanized variable region that binds to the extracellular domain of a MUC1 that is devoid of tandem repeats, a linker molecule, a transmembrane domain and a cytoplasmic domain. The single chain antibody fragment may bind to

**[0041]** (i) PSMGFR region of MUC1;

**[0042]** (ii) PSMGFR peptide;

**[0043]** (iii) a peptide having amino acid sequence of QFNQYKTEAASRYNLTISDVSVDVPPFSAQSGA (N-10) (SEQ ID NO:3)

**[0044]** (iv) a peptide having amino acid sequence of

**[0045]** ASRYNLTISDVSVDVPPFSAQSGA (N-19) (SEQ ID NO:4)

**[0046]** (v) a peptide having amino acid sequence of

[0047] NLTISDVSVSDVPPFSAQSGA (N-23) (SEQ ID NO:5)

[0048] (vi) a peptide having amino acid sequence of

[0049] ISDVSVSDVPPFSAQSGA (N-26) (SEQ ID NO:6)

[0050] (vii) a peptide having amino acid sequence of

[0051] SVSDVPPFSAQSGA (N-30) (SEQ ID NO:7)

[0052] (viii) a peptide having amino acid sequence of

[0053] QFNQYKTEAASRYNLTISDVSVSDVPPFSAQSGA (N-10/C-5) (SEQ ID NO:8)

[0054] (ix) a peptide having amino acid sequence of

[0055] ASRYNLTISDVSVSDVPPFSAQSGA (N-19/C-5) (SEQ ID NO:9)

[0056] (x) a peptide having amino acid sequence of

[0057] FPFSAQSGA (N-36) (SEQ ID NO:10)

[0058] In the CAR as described above, portions of any of the variable regions set forth and described above, or combination thereof may be used in the extracellular domain of the CAR. The CAR also comprises a transmembrane region and a cytoplasmic tail that comprises sequence motifs that signal immune system activation. The extracellular domain may be comprised of murine, camelid, human, non-human, or humanized single chain antibody fragments of an MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, and H11. Additional antibodies from which single chain antibody fragments may be made include but are not limited to monoclonal antibodies that are like MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, and H11 in that they have the same or very similar pattern of binding to subsets of peptides derived from the PSMGFR peptide, may not recognize a linear epitope or competitively inhibit the binding of NME1 or NME7<sub>AB</sub> to MUC1\*, or recognize a MUC1 transmembrane cleavage product produced by cleavage by MMP9 or contain CDR sequences that are at least 80% homologous to the MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, and H11 CDR consensus sequences.

[0059] In the CARs as described above, the extracellular domain may include a murine, camelid, human, non-human or humanized single chain antibody fragments of an MN-E6 scFv set forth as SEQ ID NOS: 233, 235, or 237), MN-C2 scFv (SEQ ID NOS:239, 241, or 243), or 20A10 scFv as set forth as SEQ ID NOS:1574-1575, 25E6 scFv as set forth as SEQ ID NOS:1598-1599.

**[0060]** In any of the CARs described above, the cytoplasmic tail may be comprised of one or more of signaling sequence motifs CD3-zeta, CD27, CD28, 4-1BB, OX40, CD30, CD40, ICAM-1, LFA-1, ICOS, CD2, CD5, or CD7. In any of the CARs described above, the cytoplasmic tails may include mutations that dampen signaling. Such mutations include but are not limited to Tyrosines that are mutated to inhibit phosphorylation and signaling (Salter et al, 2018). In any of the CARs described above, the ITAMs of CD3-zeta may be mutated to inhibit or dampen signaling (Feucht et al 2019). In any of the CARs described above, the CD3 of the cytoplasmic tail may comprise mutations in the ITAMs including those referred to as 1XX. In any of the CARs described above, the T cell may be engineered to overexpress c-Jun as a method to inhibit T cell exhaustion (Lynn et al 2019).

**[0061]** In any of the CARs described above, the sequence may be CAR MN-E6 CD28/CD3z (SEQ ID NOS:298); CAR MN-E6 4-1BB/CD3z (SEQ ID NOS:301); CAR MN-E6 OX40/CD3z (SEQ ID NOS:617); CAR MN-E6 CD28/4-1BB/CD3z (SEQ ID NOS:304); CAR MN-E6 CD28/OX40/CD3z (SEQ ID NOS:619); CAR MN-C2 CD3z (SEQ ID NOS:607); CAR MN-C2 CD28/CD3z (SEQ ID NOS:609); CAR MN-C2 4-1BB/CD3z (SEQ ID NOS:611 and SEQ ID NOS: 719); CAR MN-C2 OX40/CD3z (SEQ ID NOS:613); CAR MN-C2 CD28/4-1BB/CD3z (SEQ ID NOS: 307); CAR MN-C2 CD28/OX40/CD3z (SEQ ID NOS:615) or CAR MN-C3 4-1BB/CD3z (SEQ ID NOS: 601).

**[0062]** In another aspect, the invention is directed to a composition that includes at least two CARs with different extracellular domain units transfected into the same cell, which may be an immune cell, which may be derived from the patient requiring treatment for a cancer. The expression of the second CAR may be inducible and driven by the recognition of a target by the first CAR. The nucleic acid encoding the second CAR may be linked to an inducible promoter. The expression of the second CAR may be induced by an event that occurs specifically when the immune cell mounts an immune response to a target tumor cell. The antibody fragments of one or both of the CARs may direct the cell to a MUC1\* positive tumor. The antibody fragments of the first and second CARs may bind to a MUC1\* that is produced when MUC1 is cleaved by two different cleavage enzymes. Expression of the second CAR by the inducible promoter may be induced when the antibody fragment of the first CAR engages or binds to a MUC1 or MUC1\* on the tumor. One way to do this is to induce expression of the second CAR when, or shortly after, an NFAT protein is expressed or translocated to the nucleus. For example, a sequence



derived from an NFAT promoter region is put upstream of the gene for the second CAR. In this way, when the transcription factors that bind to the promoter of the NFAT protein are present in sufficient concentration to bind to and induce transcription of the NFAT protein, they will also bind to that same promoter that is engineered in front of the sequence for transcription of the second CAR. The NFAT protein may be NFAT1 also known as NFATc2, NFAT2 also known as NFATc or NFATc1, NFAT3 also known as NFATc4, NFAT4 also known as NFATc3, or NFAT5. In one aspect of the invention, the NFAT is NFATc1, NFATc3 or NFATc2. In one aspect of the invention, the NFAT is NFAT2 also known as NFATc1. SEQ ID NO:646 shows nucleic acid sequence of the upstream transcriptional regulatory region for NFAT2. The recognition unit of the second CAR may be an antibody fragment or a peptide, wherein the recognition units may bind to NME7, PD-1, PDL-1, or a checkpoint inhibitor.

**[0063]** The at least two CARs may have one CAR that does not have a tumor antigen targeting recognition unit and the other CAR does have a tumor antigen targeting recognition unit. In another aspect of the invention, one of the extracellular domain recognition units may bind to MUC1\* extracellular domain. In another aspect of the invention, one of the extracellular domain recognition units may be an antibody fragment and the other is a peptide, which may be devoid of transmembrane and signaling motifs; the peptide may be a single chain antibody fragment or antibody. In another aspect of the invention, one of the recognition units may bind PD-1 or PDL-1. In another aspect of the invention, one extra cellular domain recognition unit is an anti-MUC1\* antibody, antibody fragment or scFv chosen from the group consisting of MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, and H11. The other recognition unit may be a CAR or may be an anti-NME7 antibody.

**[0064]** In another aspect, the invention is directed to a cell comprising a CAR with an extracellular domain that binds to the extra cellular domain of a MUC1 molecule that is devoid of tandem repeats. In another aspect, the invention is directed to a cell comprising a CAR with an extracellular domain that binds to a MUC1\* transfected or transduced cell. The cell that includes the CAR may be an immune system cell, preferably a T cell, a natural killer cell (NK), a dendritic cell or mast cell.

**[0065]** In another aspect, the invention is directed to an engineered antibody-like protein.

**[0066]** In another aspect, the invention is directed to a method for treating a disease in a subject comprising administering an antibody according to any claim above, to a person

suffering from the disease, wherein the subject expresses MUC1 aberrantly. The disease may be cancer, such as breast cancer, ovarian cancer, pancreatic cancer, lung cancer, colon cancer, gastric cancer or esophageal cancer.

**[0067]** In another aspect, the invention is directed to an antibody, antibody fragment or scFv comprising variable domain fragments derived from an antibody that binds to an extracellular domain of MUC1 isoform or cleavage product that is devoid of the tandem repeat domains. In a preferred embodiment, the antibody or antibody fragment binds to the N-10 peptide. The variable domain fragments may be derived from mouse monoclonal antibody MN-E6 (SEQ ID NO:13 and 66) or from the humanized MN-E6 (SEQ ID NO: 39 and 94), or from MN-E6 scFv (SEQ ID NO: 233, 235 and 237). Or, the variable domain fragments may be derived from mouse monoclonal antibody MN-C2 (SEQ ID NO: 119 and 169) or from the humanized MN-C2 (SEQ ID NO: 145 and 195), or from MN-C2 scFv (SEQ ID NO: 239, 241 and 243). Or, the variable domain may be derived from monoclonal antibodies MN-18G12, MN-20A10, MN-25E6, MN-28F9, MN-5C6F3, MN-3C2B1, or MN-1E4. The heavy chain and light chain complementary determining region sequences for these antibodies are also set forth in the sequence listing herein.

**[0068]** In another aspect, the invention is directed to a method for the treatment of a person diagnosed with, suspected of having or at risk of developing a MUC1 or MUC1\* positive cancer involving administering to the person an effective amount of the antibody, antibody fragment or scFv described above, wherein the species may be murine, camelid, human or humanized.

**[0069]** In another aspect, the invention is directed to a polypeptide comprising at least two different scFv sequences, wherein one of the scFv sequences is a sequence that binds to extracellular domain of MUC1 isoform or cleavage product that is devoid of the tandem repeat domains. The polypeptide may bind to

**[0070]** (i) PSMGFR region of MUC1;

**[0071]** (ii) PSMGFR peptide;

**[0072]** (iii) a peptide having amino acid sequence of QFNQYKTEAASRYNLTISDVSVDVPPFSAQSGA (N-10) (SEQ ID NO:3)

**[0073]** (iv) a peptide having amino acid sequence of

**[0074]** ASRYNLTISDVSVDVPPFSAQSGA (N-19) (SEQ ID NO:4)

**[0075]** (v) a peptide having amino acid sequence of

[0076] NLTISDVSVSDVPPFSAQSGA (N-23) (SEQ ID NO:5)

[0077] (vi) a peptide having amino acid sequence of

[0078] ISDVSVSDVPPFSAQSGA (N-26) (SEQ ID NO:6)

[0079] (vii) a peptide having amino acid sequence of

[0080] SVSDVPPFSAQSGA (N-30) (SEQ ID NO:7)

[0081] (viii) a peptide having amino acid sequence of

[0082] QFNQYKTEAASRYNLTISDVSVSDVPPFSA (N-10/C-5) (SEQ ID NO:8)

[0083] (ix) a peptide having amino acid sequence of

[0084] ASRYNLTISDVSVSDVPPFSA (N-19/C-5) (SEQ ID NO:9)

[0085] (x) a peptide having amino acid sequence of

[0086] FPFSAQSGA (N-36) (SEQ ID NO:10)

[0087] The polypeptide may bind to a receptor on an immune cell, such as T cell, and in particular, CD3 on T-cell.

[0088] In another aspect, the invention is directed to a method of detecting presence of a cell that expresses MUC1\* aberrantly, comprising contacting a sample of cells with the scFv-Fc described above and detecting for the presence of the binding of scFv-Fc to the cell. The cell may be cancer cell.

[0089] In another aspect, the invention is directed to a method for testing a subject's cancer for suitability of treatment with a composition comprising antibodies of the invention, which may be murine, camelid, human or humanized, or fragments thereof, or portions of the variable regions of antibodies MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11, comprising the steps of contacting a bodily specimen from the patient, in vitro, ex-vivo, or in vivo, with the antibody and determining that the patient exhibits aberrant expression of MUC1\* compared to normal tissue or specimen. The antibody used in these diagnostics may be conjugated to an imaging agent.

[0090] In another aspect, the invention is directed to a method of treating a subject suffering from a disease comprising, exposing T cells from the subject, or from a donor, to MUC1\* peptides wherein through various rounds of maturation, T cells develop MUC1\* specific receptors, creating adapted T cells, and expanding and administering the adapted T cells to the donor patient who is diagnosed with, suspected of having, or is at risk of developing a MUC1\* positive cancer. The MUC1\* peptide is chosen from among the group:

- [0091] (i) PSMGFR region of MUC1;
- [0092] (ii) PSMGFR peptide;
- [0093] (iii) a peptide having amino acid sequence of QFNQYKTEAASRYNLTISDVSVDVPPFSAQSGA (N-10)
- [0094] (iv) a peptide having amino acid sequence of
- [0095] ASRYNLTISDVSVDVPPFSAQSGA (N-19)
- [0096] (v) a peptide having amino acid sequence of
- [0097] NLTISDVSVDVPPFSAQSGA (N-23)
- [0098] (vi) a peptide having amino acid sequence of
- [0099] ISDVSVDVPPFSAQSGA (N-26)
- [00100] (vii) a peptide having amino acid sequence of
- [00101] SVSDVPPFSAQSGA (N-30)
- [00102] (viii) a peptide having amino acid sequence of
- [00103] QFNQYKTEAASRYNLTISDVSVDVPPFSAQSGA (N-10/C-5)
- [00104] (ix) a peptide having amino acid sequence of
- [00105] ASRYNLTISDVSVDVPPFSAQSGA (N-19/C-5)
- [00106] (x) a peptide having amino acid sequence of
- [00107] FPFSAQSGA (N-36)
- [00108] In one aspect of the invention, the antibody that is administered to a patient for the treatment or prevention of a MUC1 or MUC1\* positive cancer is selected for its ability to bind to the N-10 peptide of the PSMGFR. The antibody can be administered alone, as a monovalent antibody, as an scFv, or a fragment of the antibody can be incorporated into a CAR, a BiTE or an ADC.
- [00109] In one aspect of the invention, the antibody that is administered to a patient for the treatment or prevention of a MUC1 or MUC1\* positive cancer is selected for its inability to recognize a linear epitope of MUC1 or MUC1\*. The antibody can be administered alone, as a monovalent antibody, as an scFv, or a fragment of the antibody can be incorporated into a CAR, a BiTE or an ADC.
- [00110] In one aspect of the invention, the antibody that is administered to a patient for the treatment or prevention of a MUC1 or MUC1\* positive cancer is selected for its ability to recognize the MUC1 transmembrane cleavage product after it has been cleaved by MMP9. The

antibody can be administered alone, as a monovalent antibody, as an scFv, or a fragment of the antibody can be incorporated into a CAR, a BiTE or an ADC.

**[00111]** In one aspect of the invention, the antibody that is administered to a patient for the treatment or prevention of a MUC1 or MUC1\* positive cancer is selected for its ability to competitively inhibit the binding of NME7<sub>AB</sub> or NME7-X1 to the extra cellular domain of a MUC1 that is devoid of tandem repeats. The antibody can be administered alone, as a monovalent antibody, as an scFv, or a fragment of the antibody can be incorporated into a CAR, a BiTE or an ADC.

**[00112]** In another aspect, the invention is directed to a method of treating cancer in a patient comprising administering to the patient the immune cell of any of the above, in combination with a checkpoint inhibitor.

**[00113]** In the method above, any of the antibodies, or variable regions thereof, set forth in the following may be used: MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11.

**[00114]** In the method above, any of the variable regions set forth in the following may be used:

**[00115]** (i) an anti-MUC1\* extracellular domain antibody or anti-N-10 antibody comprised of sequences of a humanized MN-E6 represented by humanized IgG2 heavy chain, or humanized IgG1 heavy chain, paired with humanized Kappa light chain, or humanized Lambda light chain;

**[00116]** (ii) an antibody of (i), wherein the humanized IgG2 heavy chain is SEQ ID NOS:53, humanized IgG1 heavy chain is SEQ ID NO:57, humanized Kappa light chain is SEQ ID NO:108, and humanized Lambda light chain is SEQ ID NO:112, or a sequence having 90%, 95% or 98% sequence identity thereof;

**[00117]** (iii) an anti-MUC1\* extracellular domain antibody or anti-N-10 antibody comprised of sequences of a humanized MN-C2 represented by humanized IgG1 heavy chain, humanized IgG2 heavy chain, paired with humanized Lambda light chain, and humanized Kappa light chain;

**[00118]** (iv) an antibody of (iii), wherein the humanized IgG1 heavy chain MN-C2 (SEQ ID NOS:159) or IgG2 heavy chain (SEQ ID NOS:164) paired with Lambda light chain (SEQ ID NO:219) or Kappa light chain (SEQ ID NO:213), or a sequence having 90%, 95% or 98% sequence identity thereof;

**[00119]** In the method above, in the CAR, the extracellular domain may be comprised of humanized single chain antibody fragments of MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11. The extracellular domain may be comprised of humanized single chain antibody fragments of an MN-E6 scFv set forth as SEQ ID NOS: 233, 235, or 237), MN-C2 scFv (SEQ ID NOS:239, 241, or 243). In the CAR, the cytoplasmic tail may be comprised of one or more of signaling sequence motifs CD3-zeta, CD27, CD28, 4-1BB, OX40, CD30, CD40, ICAM-1, LFA-1, ICOS, CD2, CD5, or CD7.

**[00120]** The method above may include at least two CARs with different extracellular domain units transfected into the same cell. One of the extracellular domain recognition units may bind to MUC1\* extracellular domain. One of the extracellular domain recognition units may bind to PD-1. One of the extracellular domain recognition units may be an antibody fragment and the other may be a peptide or an anti-MUC1\* antibody fragment.

**[00121]** The method may include an immune cell transfected or transduced with a plasmid encoding a CAR and a plasmid encoding a non-CAR species that is expressed from an inducible promoter. The non-CAR species may be expressed from an inducible promoter that is activated by elements of an activated immune cell. The non-CAR species may be expressed from an NFAT inducible promoter. The NFAT may be NFATc1, NFATc3 or NFATc2. The cleavage enzyme may be MMP2, MMP3, MMP9, MMP13, MMP14, MMP16, ADAM10, ADAM17, or ADAM28, or a catalytically active fragment thereof. The non-CAR species may be a cytokine. The cytokine may be IL-7, IL-12, IL-15 or IL-18.

**[00122]** The present invention is directed to an antibody, or fragment thereof, for the diagnosis, treatment or prevention of cancers wherein the antibody specifically binds to the PSMGFR peptide (SEQ ID NO:2) or a fragment thereof of the peptide.

**[00123]** The antibody binds to the N-10 peptide (SEQ ID NO:3), N-19 peptide (SEQ ID NO:4), N-23 peptide (SEQ ID NO:5), N-26 peptide (SEQ ID NO:6), N-30 peptide (SEQ ID NO:7), N-10/C-5 peptide (SEQ ID NO:8), N-19/C-5 peptide (SEQ ID NO:9), or C-5 peptide (SEQ ID NO:825).

**[00124]** The antibody interacts with a peptide comprising conformational epitope SVSDV (SEQ ID NO:1751) and FPSA (SEQ ID NO:1747) within N-26 sequence ISDVSVSDVPPFSAQSGA (SEQ ID NO:6), wherein mutation or deletion of FPFS (SEQ ID NO:1747) destroys binding of the antibody or fragment thereof to the N-26 peptide.

**[00125]** The antibody interacts with a peptide comprising conformational epitope ASRYNLT (SEQ ID NO:1745), SVSDV (SEQ ID NO:1751), and FPSA (SEQ ID NO:1747) within the N-19 sequence

ASRYNLT ISDVSVDVPPFSAQSGA (SEQ ID NO:4), wherein mutation or deletion of ASRYNLT (SEQ ID NO:1745) destroys binding of the antibody or fragment thereof to the N-26 peptide.

**[00126]** The antibody does not bind to the C-10 peptide (SEQ ID NO:825).

**[00127]** The antibody binds to the N-10 peptide (SEQ ID NO:3), but not to the C-10 peptide (SEQ ID NO:825).

**[00128]** The antibody inhibits interaction between NME7<sub>AB</sub> and MUC1\*.

**[00129]** The antibody inhibits interaction between NME7<sub>AB</sub> and PSMGFR peptide (SEQ ID NO:2).

**[00130]** The antibody inhibits interaction between NME7<sub>AB</sub> and N-10 peptide (SEQ ID NO:3), N-19 peptide (SEQ ID NO:4), N-23 peptide (SEQ ID NO:5), N-26 peptide (SEQ ID NO:6), N-30 peptide (SEQ ID NO:7), N-10/C-5 peptide (SEQ ID NO:8), N-19/C-5 peptide (SEQ ID NO:9), or C-5 peptide (SEQ ID NO:825).

**[00131]** The antibody recognizes a MUC1 transmembrane enzymatic cleavage product.

**[00132]** In the above, the cleavage enzyme is MMP14 or MMP9 or a catalytically active fragment thereof of the enzyme.

**[00133]** The antibody binds to PSMGFR (SEQ ID NO:2) or fragment thereof in which presence of an amino acid sequence within PSMGFR (SEQ ID NO:2) induces binding of the antibody to the PSMGFR.

**[00134]** The amino acid sequence of the binding conformationally inducing peptide is present in N-10 peptide (SEQ ID NO:3).

**[00135]** The antibody does not bind to a linear form of the binding conformationally inducing peptide sequence wherein the linear form of the peptide is a denatured form.

**[00136]** The binding conformationally inducing peptide sequence is in the N-26 peptide sequence ISDVSVDVPPFSAQSGA (SEQ ID NO:6), wherein mutation or deletion of FPFS (SEQ ID NO:1747) destroys binding of the antibody or fragment thereof to the N-26 peptide.

**[00137]** The binding conformationally inducing peptide sequence is located within the N-19 sequence ASRYNLTISDVSVDVPPFSAQSGA (SEQ ID NO:4), wherein mutation or deletion of ASRYNLT (SEQ ID NO:1745) destroys binding of the antibody or fragment thereof to the N-19 peptide.

**[00138]** The binding inducing peptide sequence may be located within the N-26 sequence ISDVSVDVPPFSAQSGA (SEQ ID NO:6), wherein mutation or deletion within FPFS (SEQ ID NO:1747) destroys binding of the antibody or fragment thereof to PSMGFR.

**[00139]** The antibodies may have a consensus sequence.

**[00140]** heavy chain CDR1 comprises consensus sequence at least 90% identical to sequence: F or I at position 1, T at position 2, F at position 3, S at position 4, T, G, or R at position 5, Y at position 6, A, G or T at position 7, M at position 8 and S at position 9;

**[00141]** heavy Chain CDR2 comprises consensus sequence at least 90% identical to sequence: T at position 1, I or S at position 2, I or S at position 3, G or R at position 5, G or A at position 6, T or I at position 9, Y at position 10, Y at position 11, P or S at position 12 and DSVKG for positions 13-17;

**[00142]** heavy chain CDR3 comprises consensus sequence at least 90% identical to sequence: \_ G, L, or N at position 2, G or T at position 4, Y at position 7, D or E at position 12, A at position 14, and Y at position 15;

**[00143]** light chain CDR1 comprises consensus sequence at least 90% identical to sequence: K or R at position 1, A or S at position 2, S at position 3, K or Q at position 4, S at position 5, L or V at position 6, L at position 7, T or S at position 10, Y at position 15, and I, L or M at position 16;

**[00144]** light Chain CDR2 comprises consensus sequence at least 90% identical to sequence: L or W, or S at position 1, A or T at position 2, S at position 3, N or T at position 4, L or R at position 5, E or A at position 6, and S at position 7; and

**[00145]** light chain CDR3 comprises consensus sequence at least 90% identical to sequence: Q at position 1, H or Q at position 2, S, Q or R at position 3, R, S or Y at position 4, E, L, or S at position 5, L or S at position 6, P or S at position 7, F or L at position 8 and T at position 9.

**[00146]** An antibody binding conformationally inducing peptide is within the N-26 sequence ISDVSVSDVPFPFSAQSGA (SEQ ID NO:6), wherein mutation or deletion within FPFS (SEQ ID NO:1747), SVSDV (SEQ ID NO:1751), or ASRYNLT (SEQ ID NO:1745) destroys binding of the antibody or fragment thereof to PSMGFR.

**[00147]** The antibody may have a further consensus sequence,

**[00148]** wherein

**[00149]** heavy chain CDR1 comprises consensus sequence at least 90% identical to sequence: F or I at position 1, T or A at position 2, F at position 3, S at position 4, T, G, or R at position 5, Y or F at position 6, A, G or T at position 7, M at position 8 and S at position 9;

**[00150]** heavy Chain CDR2 comprises consensus sequence at least 90% identical to sequence: T or A at position 1, I or S at position 2, I or S at position 3, N, S, T or G at position 4, G or R at position 5, G or A at position 6, G, T, or D at position 7, Y, K, H or S at position 8, T or I at position 9, Y or F at position 10, Y at position 11, P or S at position 12 and D at position 13, S or T at position 14, V or L at position 15 and KG for positions 16-17;

**[00151]** heavy chain CDR3 comprises consensus sequence at least 90% identical to sequence: G, L, or N at position 2, G, T, or Y at position 3, G or T at position 4, Y at position 7, Y, A, or G at position 10, M, D or F at position 11, D or E at position 12 and AY at position 14-15;



- [00152]** light chain CDR1 comprises consensus sequence \_ at least 90% identical to sequence: K or R at position 1, A or S at position 2, S or R at position 3, S, Y, I or V at position 8, T or S at position 10, G, S, D, or Q at position 12, V, Y, K or N at position 13, N, S, or T at position 14, Y or F at position 15, and I, L or M at position 16;
- [00153]** light Chain CDR2 comprises consensus sequence at least 90% identical to sequence: A, T or V at position 2, S at position 3, N, T, or K at position 4, L or R at position 5, E, A, F or D at position 6, and S at position 7; and
- [00154]** light chain CDR3 comprises consensus sequence at least 90% identical to sequence: Q, F or W at position 1, H or Q at position 2, R, S, T, Y or N at position 4, E, L, S or H at position 5, L, S, V, D or Y at position 6, P or S at position 7, and T at position 9.
- [00155]** The antibody above which may be MNC2, having
- [00156]** heavy chain CDR1 comprises consensus sequence FTFSGYAMS;
- [00157]** heavy Chain CDR2 comprises consensus sequence TISSGGTYIYYPDSVKG;
- [00158]** heavy chain CDR3 comprises consensus sequence -LGGDNYEYFDV--;
- [00159]** light chain CDR1 comprises consensus sequence RASKS--VSTSGYSYMH;
- [00160]** light Chain CDR2 comprises consensus sequence LASNLES; and
- [00161]** light chain CDR3 comprises consensus sequence QHSRELPFT.
- [00162]** MNE6, having
- [00163]** heavy chain CDR1 comprises consensus sequence FTFSRYGMS;
- [00164]** heavy Chain CDR2 comprises consensus sequence TISGGGTYYIYPDSVKG;
- [00165]** heavy chain CDR3 comprises consensus sequence DNYGRNYDYGMDY--;
- [00166]** light chain CDR1 comprises consensus sequence -----SATSSVSYIH;
- [00167]** light Chain CDR2 comprises consensus sequence STSNLAS; and
- [00168]** light chain CDR3 comprises consensus sequence QQRSSSPFT.
- [00169]** B2, having
- [00170]** heavy chain CDR1 comprises consensus sequence FAFSTFAMS;
- [00171]** heavy Chain CDR2 comprises consensus sequence AISNGGGYTYYPDTLKG;
- [00172]** heavy chain CDR3 comprises consensus sequence ----RYYDLYFDL--;
- [00173]** light chain CDR1 comprises consensus sequence RSSQNIV-HSNGNTYLE;
- [00174]** light Chain CDR2 comprises consensus sequence KVSNRFS; and
- [00175]** light chain CDR3 comprises consensus sequence FQDSHVPLT.
- [00176]** B7, having
- [00177]** heavy chain CDR1 comprises consensus sequence FTFSRYGMS;

- [00178] heavy Chain CDR2 comprises consensus sequence TISSGGTYIYYPDSVKG;
- [00179] heavy chain CDR3 comprises consensus sequence DNYGSSYDYAMDY--;
- [00180] light chain CDR1 comprises consensus sequence RSSQTIV-HSNGNTYLE;
- [00181] light Chain CDR2 comprises consensus sequence KVSNRFS; and
- [00182] light chain CDR3 comprises consensus sequence FQDSHVPLT.
- [00183] B9, having
- [00184] heavy chain CDR1 comprises consensus sequence FTFSRYGMS;
- [00185] heavy Chain CDR2 comprises consensus sequence TISSGGTYIYYPDSVKG;
- [00186] heavy chain CDR3 comprises consensus sequence DNYGSSYDYAMDY--;
- [00187] light chain CDR1 comprises consensus sequence -----SASSSVSYM;
- [00188] light Chain CDR2 comprises consensus sequence TTSNLA; and
- [00189] light chain CDR3 comprises consensus sequence QQRSSYPF-.
- [00190] 8C7F3, having
- [00191] heavy chain CDR1 comprises consensus sequence FTFSTYAMS;
- [00192] heavy Chain CDR2 comprises consensus sequence AISNGGGYTYYPDSLKG;
- [00193] heavy chain CDR3 comprises consensus sequence ----RYYDHYFDY--;
- [00194] light chain CDR1 comprises consensus sequence --RASESVATYGNNFMQ;
- [00195] light Chain CDR2 comprises consensus sequence LASTLDS; and
- [00196] light chain CDR3 comprises consensus sequence QQNNEDPPT.
- [00197] H11, having
- [00198] heavy chain CDR1 comprises consensus sequence FAFSTFAMS;
- [00199] heavy Chain CDR2 comprises consensus sequence AISNGGGYTYYPDTLKG;
- [00200] heavy chain CDR3 comprises consensus sequence ----RYYDLYFDL--;
- [00201] light chain CDR1 comprises consensus sequence RSSQNIV-HSNGNTYLE;
- [00202] light Chain CDR2 comprises consensus sequence KVSNRFS; and
- [00203] light chain CDR3 comprises consensus sequence FQDSHVPLT.
- [00204] B12, having
- [00205] heavy chain CDR1 comprises consensus sequence SYGVH;
- [00206] heavy Chain CDR2 comprises consensus sequence VIWPGGSTNYNSTLMSRM;
- [00207] heavy chain CDR3 comprises consensus sequence DRTPRVGAWFAY; and
- [00208] light chain CDR1 comprises consensus sequence RASESVATYGNNFMQ;
- [00209] light Chain CDR2 comprises consensus sequence LASTLDS; and

- [00210] light chain CDR3 comprises consensus sequence QQNNEDPPT.
- [00211] 20A10, having
- [00212] heavy chain CDR1 comprises consensus sequence FTFSTYAMS;
- [00213] heavy Chain CDR2 comprises consensus sequence -SIGRAGSTYYSDSVKG;
- [00214] heavy chain CDR3 comprises consensus sequence ---GPIYNDYDEFAY;
- [00215] light chain CDR1 comprises consensus sequence KSSQSVLYSSNQKNYLA;
- [00216] light Chain CDR2 comprises consensus sequence WASTRES; and
- [00217] light chain CDR3 comprises consensus sequence HQYLSSLT.
- [00218] 3C2B1, having
- [00219] heavy chain CDR1 comprises consensus sequence ITFSTYTMS;
- [00220] heavy Chain CDR2 comprises consensus sequence TISTGGDKTYYSDSVKG;
- [00221] heavy chain CDR3 comprises consensus sequence -GTTAMYYYAMDY;
- [00222] light chain CDR1 comprises consensus sequence RASKS---ISTSDYNYIH ;
- [00223] light Chain CDR2 comprises consensus sequence LASNLES; and
- [00224] light chain CDR3 comprises consensus sequence QHSRELPLT.
- [00225]
- [00226] In another aspect, the invention is directed to an antibody, or fragment thereof, for the diagnosis, treatment or prevention of cancers that requires presence of antibody binding conformationally inducing peptide ASRYNLT (SEQ ID NO:1745) of PSMGFR (SEQ ID NO:2). The antibody may be 25E6, having
- [00227] heavy chain CDR1 comprises consensus sequence FTFSSYGMS;
- [00228] heavy Chain CDR2 comprises consensus sequence TISNGGRHTFYPPDSVKG;
- [00229] heavy chain CDR3 comprises consensus sequence QTGTEGWFAFAY;
- [00230] light chain CDR1 comprises consensus sequence KSSQSLLDSDGKTYLN;
- [00231] light Chain CDR2 comprises consensus sequence LVSKLDS \_; and
- [00232] light chain CDR3 comprises consensus sequence WQGTHFPQT.
- [00233]
- [00234] In another aspect, the invention is directed to an antibody, or fragment thereof, for the diagnosis, treatment or prevention of cancers that requires presence of antibody binding conformationally inducing peptide SVSDV (SEQ ID NO:1761) of PSMGFR (SEQ ID NO:2). The antibody may be 5C6F3, having
- [00235] heavy chain CDR1 comprises consensus sequence FTFSTYAMS ;
- [00236] heavy Chain CDR2 comprises consensus sequence AISNGGGYTYYPDSLKG;

**[00237]** heavy chain CDR3 comprises consensus sequence RYYDHYFDY;

**[00238]** light chain CDR1 comprises consensus sequence RSSQTIVHSNGNTYLE;

**[00239]** light Chain CDR2 comprises consensus sequence KVSNRFS; and

**[00240]** light chain CDR3 comprises consensus sequence FQDSHVPLT.

**[00241]** The antibody or fragment thereof according all of the above may be murine, camelid, human or humanized. The antibody fragment may be scFv or scFv-Fc, which variable regions thereof may be murine, camelid, human or humanized.

**[00242]** In another aspect, the invention is directed to a chimeric antigen receptor (CAR) comprising the antibody fragments of above, and may further comprise mutations in the co-stimulatory domain or CD3-zeta signaling domain. Tyrosines may be mutated in CD28 or 4-1BB. CD3-zeta may contain 1XX mutations.

**[00243]** In another aspect, the invention is directed to an immune cell comprising the CAR of above. Immune cell may be T cell, NK cell, dendritic cell, or mast cell.

**[00244]** In another aspect, the invention is directed to a cell composition expressed in a cell comprising a CARs of above, and second entity having a biological recognition unit that has a specificity that is different from that of the CAR. The second entity may bind PD-1, PDL-1, or other checkpoint inhibitor, or NME7, or a cytokine such as IL-12 or IL-18, or c-Jun.

**[00245]** In yet another aspect, the invention is directed to an immune cell engineered to express a nucleic encoding a CAR of above and a nucleic acid encoding a second entity as in any of the claims above wherein the second entity expressed from an inducible promoter. The second entity may be expressed from an inducible promoter that is activated by elements of an activated immune cell. The second entity may be expressed from an NFAT inducible promoter. NFAT may be NFATc1, NFATc3 or NFATc2. The second entity may be a cytokine such as IL-7, IL-15, or IL-18. The nucleic acids encoding the second entity may be inserted into a Foxp3 promoter or enhancer region, wherein the cytokine is IL-18. The cytokine may be expressed from an NFAT inducible promoter.

**[00246]** In another aspect, the invention is directed to a BiTE construct comprising the antibody fragment of above.

**[00247]** In yet another aspect, the invention is directed to an antibody drug conjugate (ADC) comprising the antibody or antibody fragment of above.

- [00248] The present invention is directed to an antibody or fragment thereof that specifically binds to PSMGFR (SEQ ID NO:2) and N-10 (SEQ ID NO:3); and
- [00249] does not bind to full-length MUC1;
- [00250] does not bind to C-10 (SEQ ID NO:825);
- [00251] competitively inhibits binding of NME1 or NME7<sub>AB</sub> to MUC1\* extra cellular domain or a PSMGFR peptide;
- [00252] recognizes a MUC1\* generated by cleavage by a cleavage enzyme;
- [00253] recognizes a conformational epitope and not a linear epitope; or
- [00254] is cancer selective by immunohistochemistry on tissues.
- [00255] Four of the criteria (i) – (vi) may be satisfied. Five of the criteria (i) – (vi) may be satisfied. Six of the criteria (i) – (vi) may be satisfied. At least criteria (vi) may be satisfied. Cleavage enzyme may be MMP-9.
- [00256] In all of the above, the cancer may be breast cancer, pancreatic cancer, ovarian cancer, lung cancer, colon cancer, gastric cancer or esophageal cancer.
- [00257] The present invention is also directed to a method of diagnosing, treating or preventing cancer by administering the antibodies and fragments disclosed herein to a cancer patient in need thereof that has been identified as expressing MUC1 aberrantly and expressing truncated MUC1, such as MUC1\*.
- [00258] These and other objects of the invention will be more fully understood from the following description of the invention, the referenced drawings attached hereto and the claims appended hereto.

## **BRIEF DESCRIPTION OF THE DRAWINGS**

- [00259] The patent or application file contains at least one drawing executed in color. Copies of this patent or patent application publication with color drawings will be provided by the Office upon request and payment of the necessary fee.
- [00260] The present invention will become more fully understood from the detailed description given herein below, and the accompanying drawings which are given by way of illustration only, and thus are not limitative of the present invention, and wherein;
- [00261] **Figures 1A-1D** show cell growth assay graphs of MUC1\* positive cells treated with either bivalent ‘bv’ anti-MUC1\* antibody, monovalent ‘mv’ or Fab, NM23-H1 dimers or

NME7-AB. Bivalent anti-MUC1\* antibodies stimulate growth of cancer cells whereas the monovalent Fab inhibits growth (Fig. 1A-1B). Classic bell-shaped curve indicates ligand induced dimerization stimulates growth. Dimeric NM23-H1, aka NME1, stimulates growth of MUC1\* positive cancer cells but siRNA to suppress MUC1 expression eliminate its effect (Fig. 1C). NME7-AB also stimulates the growth of MUC1\* positive cells (Fig. 1D).

**[00262]** **Figures 2A-2I** show results of ELISA assays. MUC1\* peptides PSMGFR, PSMGFR minus 10 amino acids from the N-terminus aka N-10, or PSMGFR minus 10 amino acids from the C-terminus, aka C-10 are immobilized on the plate and the following are assayed for binding: NME7-AB (Fig. 2A), MN-C2 monoclonal antibody (Fig. 2B), MN-E6 monoclonal antibody (Fig. 2C), or dimeric NME1 (Fig. 2D). These assays show that NME1, NME7-AB and monoclonal antibodies MN-C2 and MN-E6 all require the first membrane proximal 10 amino acids of the MUC1\* extracellular domain to bind. MUC1\* peptides PSMGFR minus 10 amino acids from the N-terminus aka N-10, or PSMGFR minus 10 amino acids from the C-terminus, aka C-10, are immobilized on the plate and the following are assayed for binding: MN-C3 (Fig. 2E) and MN-C8 (Fig. 2F). Fig. 2G shows the amino acid sequence of the PSMGFR peptide. Fig. 2H shows the amino acid sequence of the N-10 peptide. Fig. 2I shows the amino acid sequence of the C-10 peptide.

**[00263]** **Figures 3A-3C** show results of competitive ELISA assays. The PSMGFR MUC1\* peptide is immobilized on the plate and dimeric NM23-H1, aka NME1, is added either alone or after the MN-E6 antibody has been added (Fig. 3A). The same experiment was performed wherein NM23-H7, NME7-AB, is added alone or after MN-E6 has been added (Fig. 3B). Results show that MN-E6 competitively inhibits the binding of MUC1\* activating ligands NME1 and NME7. In a similar experiment (Fig. 3C), PSMGFR or PSMGFR minus 10 amino acids from the N-terminus, aka N-10, is immobilized on the plate. Dimeric NM23-H1 is then added. Anti-MUC1\* antibodies MN-E6, MN-C2, MN-C3 or MN-C8 are then tested for their ability to compete off the NM23-H1. Results show that although all three antibodies bind to the PSMGFR peptides, MN-E6 and MN-C2 competitively inhibit binding of the MUC1\* activating ligands.

**[00264]** **Figures 4A-4F** show FACS scans of anti-MUC1\* antibody huMN-C2scFv binding specifically to MUC1\* positive cancer cells and MUC1\* transfected cells but not MUC1\* or MUC1 negative cells. ZR-75-1, aka 1500, MUC1\* positive breast cancer cells were stained with 1:2 or 1:10 dilutions of the 1.5 ug/ml humanized MN-C2. After two washes, cells were stained

with secondary antibody, Anti-Penta-His antibody conjugated to Alexa 488 (Qiagen) dilutions of 1:200 (Fig. 4A), 1:50 (Fig. 4B), or 1:10 (Fig. 4C) to detect the 6x His tag on the huMN-C2 scFv. Fig. 4A shows huMN-C2 binding to ZR-75-1 breast cancer cells where secondary antibody is added at a 1:200 dilution. Fig. 4B shows huMN-C2 binding to ZR-75-1 breast cancer cells where secondary antibody is added at a 1:50 dilution. Fig. 4C shows huMN-C2 binding to ZR-75-1 breast cancer cells where secondary antibody is added at a 1:10 dilution. Flow cytometric analysis revealed a concentration-dependent shift of a subset of cells, indicating specific binding, which is unseen in the absence of the MN-C2 scFv (Fig. 4A-4C). Fig. 4D shows anti-MUC1\* antibody MN-E6 staining of MUC1 negative HCT-116 colon cancer cells transfected with the empty vector, single cell clone #8. Fig. 4E shows anti-MUC1\* antibody MN-E6 staining of HCT-116 colon cancer cells transfected with MUC1\* single cell clone #10. Fig. 4F shows anti-MUC1\* antibody MN-E6 staining of ZR-75-1, aka 1500, MUC1\* positive breast cancer cells. As the FACS scans show, both MN-C2 and MN-E6 only stain MUC1\* positive cells and not MUC1 or MUC1\* negative cells.

**[00265]** **Figure 5** shows a graph of an ELISA in which surface is coated with either the MUC1\* PSMGFR peptide or a control peptide. Humanized MN-C2 scFv is then incubated with the surface, washed and detected according to standard methods. The ELISA shows that the huMN-C2 scFv binds to the MUC1\* peptide with an EC-50 of about 333nM.

**[00266]** **Figures 6A-6B** show graphs of cancer cell growth inhibition by MUC1\* antibody variable region fragment humanized MN-C2 scFv. hMN-C2 scFv potently inhibited the growth of ZR-75-1, aka 1500, MUC1\* positive breast cancer cells (Fig. 6A) and T47D MUC1\* positive breast cancer cells (Fig. 6B) with approximately the same EC-50 as the *in vitro* ELISAs.

**[00267]** **Figures 7A-7B** show graphs of tumor growth in immune compromised mice that have been implanted with human tumors then treated with anti-MUC1\* antibody MN-E6 Fab or mock treatment. Female nu/nu mice implanted with 90-day estrogen pellets were implanted with 6 million T47D human breast cancer cells that had been mixed 50/50 with Matrigel. Mice bearing tumors that were at least 150 mm<sup>3</sup> and had three successive increases in tumor volume were selected for treatment. Animals were injected sub cutaneously twice per week with 80 mg/kg MN-E6 Fab and an equal number of mice fitting the same selection criteria were injected with vehicle alone (Fig. 7A). Male NOD/SCID mice were implanted with 6 million DU-145 human prostate cancer cells that had been mixed 50/50 with Matrigel. Mice bearing tumors that

were at least 150 mm<sup>3</sup> and had three successive increases in tumor volume were selected for treatment. Animals were injected sub-cutaneously every 48 hours with 160 mg/kg MN-E6 Fab and an equal number of mice fitting the same selection criteria were injected with vehicle alone (Fig. 7B). Tumors were measured independently by two researchers twice per week and recorded. Statistics were blindly calculated by independent statistician, giving a P value of 0.0001 for each. Anti-MUC1\* Fab inhibited breast cancer growth and prostate cancer growth. Treatment had no effect on weight, bone marrow cell type or number.

**[00268]** **Figure 8** shows a graph of an ELISA wherein the surface was immobilized with either PSMGFR peptide, PSMGFR minus 10 amino acids from the N-terminus or minus 10 amino acids from the C-terminus. The huMN-E6 scFv-Fc bound to the PSMGFR peptide and to the PSMGFR N-10 peptide but not to the PSMGFR C-10 peptide. The parent MN-E6 antibody and the humanized MN-E6 require the C-terminal 10 amino acids of PSMGFR for binding.

**[00269]** **Figures 9A-9B** show graphs of ELISAs wherein the assay plate surface was immobilized with either PSMGFR peptide, PSMGFR minus 10 amino acids from the N-terminus or minus 10 amino acids from the C-terminus. The MN-C3 antibody variants were then assayed for binding to the various MUC1\* peptides. Fig. 9A shows purified mouse monoclonal MN-C3 antibody; and Figure 9B shows the humanized MN-C3 scFv-Fc. ELISAs show binding to the PSMGFR peptide as well as to certain deletion peptides.

**[00270]** **Figures 10A-10J.** Figs. 10A10B are photographs of breast cancer tissue arrays. Fig. 10A was stained with VU4H5 which recognizes MUC1-FL (full length); Fig. 10B was stained with mouse monoclonal antibody MN-C2 which recognizes cancerous MUC1\*. Following automated staining (Clariant Diagnostics), the tissue staining was scored using Allred scoring method which combines an intensity score and a distribution score. Figs. 10C10F are color coded graphs showing the score calculated for MUC1 full-length staining for each patient's tissue. Figs. 10G10J are color coded graphs showing the score calculated for MUC1\* staining for each patient's tissue.

**[00271]** **Figures 11A-11J.** Figs. 11A11B are photographs of breast cancer tissue arrays. Fig. 11A was stained with VU4H5 which recognizes MUC1-FL (full length); Fig. 11B was stained with mouse monoclonal antibody MN-C2 which recognizes cancerous MUC1\*. Following automated staining (Clariant Diagnostics), the tissue staining was scored using Allred scoring method which combines an intensity score and a distribution score. Figs. 11C-11F are color



coded graphs showing the score calculated for MUC1 full-length staining for each patient's tissue. Figs. 11G-11J are color coded graphs showing the score calculated for MUC1\* staining for each patient's tissue.

**[00272]** **Figures 12A-12H** show photographs of normal breast and breast cancer tissues stained with humanized MN-E6-scFv-Fc biotinylated anti-MUC1\* antibody at 2.5 ug/mL, then stained with a secondary streptavidin HRP antibody. Fig. 12A is a normal breast tissue. Figs. 12B-12D are breast cancer tissues from patients as denoted in the figure. Figs. 12E-12H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00273]** **Figures 13A-13F** show photographs of normal breast and breast cancer tissues stained with humanized MN-E6-scFv-Fc biotinylated anti-MUC1\* antibody at 2.5 ug/mL, then stained with a secondary streptavidin HRP antibody. Fig. 13A is a normal breast tissue. Figs. 13B-13C are breast cancer tissues from patients as denoted in the figure. Figs. 13D-13F are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00274]** **Figures 14A-14H** show photographs of breast cancer tissues stained with MN-E6 anti-MUC1\* antibody at 10 ug/mL, then stained with a rabbit anti mouse secondary HRP antibody. Figs. 14A-14D are breast cancer tissues from patient #300. Figs. 14E-14H are breast cancer tissues from metastatic patient #291.

**[00275]** **Figures 15A-15F** show photographs of normal lung and lung cancer tissues stained with humanized MN-E6-scFv-Fc biotinylated anti-MUC1\* antibody at 2.5 ug/mL, then stained with a secondary streptavidin HRP antibody. Fig. 15A is a normal lung tissue. Figs. 15B-15C are lung cancer tissues from patients as denoted in the figure. Figs. 15D-15F are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00276]** **Figures 16A-16F** show photographs of normal lung and lung cancer tissues stained with humanized MN-E6-scFv-Fc biotinylated anti-MUC1\* antibody at 2.5 ug/mL, then stained with a secondary streptavidin HRP antibody. Fig. 16A is a normal lung tissue. Figs. 16B-16C are lung cancer tissues from patients as denoted in the figure. Figs. 16D-16F are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00277]** **Figures 17A-17F** show photographs of normal lung and lung cancer tissues stained with humanized MN-E6-scFv-Fc biotinylated anti-MUC1\* antibody at 25 ug/mL, then stained

with a secondary streptavidin HRP antibody. Fig. 17A is a normal lung tissue. Figs. 17B-17C are lung cancer tissues from patients as denoted in the figure. Figs. 17D-17F are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00278]** **Figures 18A-18F** show photographs of normal lung and lung cancer tissues stained with humanized MN-E6-scFv-Fc biotinylated anti-MUC1\* antibody at 25 ug/mL, then stained with a secondary streptavidin HRP antibody. Fig. 18A is a normal lung tissue. Figs. 18B-18C are lung cancer tissues from patients as denoted in the figure. Figs. 18D-18F are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00279]** **Figures 19A-19D** show photographs of normal small intestine and cancerous small intestine tissues stained with humanized MN-E6-scFv-Fc biotinylated anti-MUC1\* antibody at 5 ug/mL, then stained with a secondary streptavidin HRP antibody. Fig. 19A is a normal small intestine tissue. Fig. 19B is small intestine cancer from patient as denoted in the figure. Figs. 19C-19D are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00280]** **Figures 20A-20H** show photographs of normal small intestine tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. Figs. 20A-20D are normal small intestine tissue. Figs. 20E-20H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00281]** **Figures 21A-21H** show photographs of cancerous small intestine tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. Figs. 21A-21D are cancerous small intestine tissue from a patient as denoted in figure. Figs. 21E-21H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00282]** **Figures 22A-22H** show photographs of cancerous small intestine tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. Figs. 22A-22D are cancerous small intestine tissue from a patient as denoted in figure. Figs. 22E-22H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00283]** **Figures 23A-23H** show photographs of normal colon tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-

human HRP antibody. Figs. 23A-23D are normal colon. Figs. 23E-23H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00284] Figures 24A-24H** show photographs of colon cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. Figs. 24A-24D are colon cancer tissue from a metastatic patient as denoted in figure. Figs. 24E-24H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00285] Figures 25A-25H** show photographs of colon cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. Figs. 25A-25D are colon cancer tissue from a Grade 2 patient as denoted in figure. Figs. 25E-25H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00286] Figures 26A-26H** show photographs of colon cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. Figs. 26A-26D are colon cancer tissue from a metastatic patient as denoted in figure. Figs. 26E-26H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00287] Figures 27A-27H** show photographs of prostate cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. Figs. 27A-27D are prostate cancer tissue from a patient as denoted in figure. Figs. 27E-27H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00288] Figures 28A-28H** show photographs of prostate cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. Figs. 28A-28D are prostate cancer tissue from a patient as denoted in figure. Figs. 28E-28H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00289] Figures 29A-29H** show photographs of prostate cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. Figs. 29A-29D are prostate cancer tissue from a patient as

denoted in figure. Figs. 29E-29H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00290]**

**[00291]** **Figures 30A-30F** show photographs of a triple negative breast cancer array stained with anti-MUC1\* antibody huMNC2scFv. The first score shown is the Allred score and the second is the tumor grade. The percentage of the array that scored zero, weak, medium or strong is graphed as a pie chart. Fig. 30A shows the pie chart of score of anti-MUC1\* antibody staining. Fig. 30B shows a photograph of the array stained with the antibody. Figs. 30C-30D show magnified photographs of two of the breast cancer specimens from the array. Figs. 30E-30F show more magnified photographs of the portion of the specimen that is marked by a box.

**[00292]** **Figures 31A-31F** show photographs of an ovarian cancer array stained with anti-MUC1\* antibody huMNC2scFv. The first score shown is the Allred score and the second is the tumor grade. The percentage of the array that scored zero, weak, medium or strong is graphed as a pie chart. Fig. 31A shows the pie chart of score of anti-MUC1\* antibody staining. Fig. 31B shows a photograph of the array stained with the antibody. Figs. 31C-31D show magnified photographs of two of the breast cancer specimens from the array. Figs. 31E-31F show more magnified photographs of the portion of the specimen that is marked by a box.

**[00293]** **Figures 32A-32F** show photographs of a pancreatic cancer array stained with anti-MUC1\* antibody huMNC2scFv. The first score shown is the Allred score and the second is the tumor grade. The percentage of the array that scored zero, weak, medium or strong is graphed as a pie chart. Fig. 32A shows the pie chart of score of anti-MUC1\* antibody staining. Fig. 32B shows a photograph of the array stained with the antibody. Figs. 32C-32D show magnified photographs of two of the breast cancer specimens from the array. Figs. 32E-32F show more magnified photographs of the portion of the specimen that is marked by a box.

**[00294]** **Figures 33A-33F** show photographs of a lung cancer array stained with anti-MUC1\* antibody huMNC2scFv. The first score shown is the Allred score and the second is the tumor grade. The percentage of the array that scored zero, weak, medium or strong is graphed as a pie chart. Fig. 33A shows the pie chart of score of anti-MUC1\* antibody staining. Fig. 33B shows a photograph of the array stained with the antibody. Figs. 33C-33D show magnified photographs of two of the breast cancer specimens from the array. Figs. 33E-33F show more magnified photographs of the portion of the specimen that is marked by a box.

[00295] **Figures 34A-34I** show photographs of normal tissues stained with anti-MUC1\* antibody huMNC2scFv.

[00296] **Figures 35A-35D** show FACS scans of cells expressing either no MUC1, MUC1\* or full-length MUC1, wherein the cells were probed with either MNC2 or VU4H5. Fig. 35A shows MUC1 negative HCT-116 colon cancer cells probed with antibody MNC2. Fig. 35B shows HCT cells that have been transfected with MUC1\* wherein the extra cellular domain is just the sequence of the PSMGFR peptide wherein the cells are probed with antibody MNC2. Fig. 35C shows HCT-MUC1-18 cells which are a cleavage resistant single cell clone of HCT cells transfected with full-length MUC1, also referred to herein as HCT-MUC1-41TR, and cells were probed with antibody MNC2. Fig. 35D shows HCT-MUC1-18 cells probed with antibody VU4H5 which is an antibody that recognizes the hundreds of tandem repeats epitopes in full-length MUC1. As can be seen in the figures, MNC2 recognizes an ectopic epitope that is not accessible in full-length MUC1.

[00297] **Figures 36A-36D** show Western blots and corresponding FACS analysis of HCT-116 cells which are a MUC1 negative colon cancer cell line, that were then stably transfected with either MUC1\* or MUC1 full-length. The single cell clones that are shown are HCT-MUC1-41TR, and HCT-MUC1\*. Fig. 36A shows a Western blot of the parent cell line HCT-116, HCT-MUC1-41TR and HCT-MUC1\* wherein the gel has been probed with a rabbit polyclonal antibody, SDIX, that only recognizes cleaved MUC1. A visible band between 25 and 35kDa can be readily seen in Lane 6, loaded with HCT-MUC1\*, whereas there is only a faint band in Lanes 4 and 5, showing that only a small amount of MUC1 is cleaved in the HCT-MUC1-41Tr cells. There is no cleaved MUC1 present in the parent cell line HCT-116 loaded into Lanes 2 and 3. Fig. 36B is a Western blot that was probed with a mouse monoclonal antibody VU4H5 that recognizes the tandem repeats of full-length MUC1. As can be seen, only HCT-MUC1-41TR contains full-length MUC1. Fig. 36C shows FACS scans showing that HCT-MUC1\* is 95.7% positive for SDIX which only binds to MUC1\* and essentially not at all for MUC1 full-length. Fig. 36D shows FACS scans that show that HCT-MUC1-41TR cells are 95% positive for full-length MUC1 and only about 11% positive for the cleaved form, MUC1\*.

[00298] **Figure 37A-37C** shows western blots and a bar graph of FACS analysis assessing the ability of MNC2 to recognize a full-length MUC1 after it has been cleaved by MMP9. Fig. 37A shows a Western blot of HCT-MUC1-18 cells, which are a cleavage resistant cell line, to which

was added cleavage enzyme MMP9. The cell lysate fraction was run on a gel and probed with a polyclonal anti-PSMGFR antibody. The photo shows that in a dose dependent manner, MMP9 cleaved MUC1 to MUC1\*, the ~25kDa species. Fig. 37B shows the Western blot of the conditioned media from the same experiment. The photo shows that the addition of cleavage enzyme MMP9, in a dose dependent manner, increased the release of the tandem repeat domain into the conditioned media. Fig. 37C shows FACS analysis of the experiment. The graphs show that the addition of MMP9, in a dose dependent manner, increased recognition of the cleavage product by anti-MUC1\* antibody MNC2 and decreased the recognition of the full-length MUC1 which contains the tandem repeat domain.

**[00299]** **Figure 38** shows a photograph of a Western blot in which HCT-MUC1-18 cells, labeled here as HCT-18, a cleavage resistant single cell clone of HCT cells transfected with full-length MUC1, are treated with varying amounts of a catalytically active ADAM17 or MMP14. Shed MUC1 tandem repeat domain of full-length MUC1 is immunoprecipitated from the conditioned media, and run on a gel that is then probed with VU4H5 that binds to the tandem repeat epitopes. As can be seen, MMP14 also efficiently cleaves MUC1 full-length and sheds the tandem repeat containing extra cellular domain into the conditioned media. Cleavage enzyme ADAM17 did not cleave MUC1.

**[00300]** **Figure 39A-39B** shows fluorescence activated cell sorting (FACS) measurements of human CD34+ hematopoietic stem cells of human bone marrow stained with anti-MUC1\* monoclonal antibodies MNC3, MNC2, MNE6 or an isotype control antibody. The histogram of the FACS assay and the bar graph showing the data show that the MUC1\* positive cells of the bone marrow are recognized by one anti-MUC1\* antibody, MNC3 but not by MNE6 or MNC2. All three antibodies bind to the PSMGFR peptide. The great difference in the specificity of these antibodies suggests that MNC3 recognizes a MUC1\*-like form created when MUC1 is cleaved by an enzyme that is different from MMP9.

**[00301]** **Figure 40A-40G** shows the details of FACS analysis of the hematopoietic stem cells probed with either MNC3 or MNE6. Fig. 40A shows the FACS scatter plot of total bone marrow cells. Fig. 40B shows the FACS scatter plot of the CD34+ cells. Fig. 40C shows the FACS histogram of the CD34+ cells. Fig. 40D shows the FACS scatter plot of the earliest hematopoietic stem cells, which are CD34+/CD38-, stained with either MNC3 or MNE6. Fig. 40E shows the histogram of the experiment. Fig. 40F shows the histogram overlay of MNC3

binding to CD34+/CD38<sup>-</sup> cells versus MNE6. Fig. 40G shows the bar graph of that FACS experiment.

**[00302]** **Figure 41A-42H** shows the details of FACS analysis of CD34+/CD38<sup>-/lo</sup> hematopoietic stem cells probed with a polyclonal anti-PSMGFR antibody SDIX, MNE6 or MNC2. Fig. 41A shows the FACS scatter plot of the CD34+/CD38<sup>-/lo</sup> population of cells. Fig. 41E shows a table of the detailed analysis. Fig. 41B shows the FACS scatter plot of the CD34+/CD38<sup>-/lo</sup> population of cells probed with the anti-PSMGFR polyclonal antibody SDIX. Fig. 41F shows a table of the detailed analysis. Fig. 41C shows the FACS scatter plot of the CD34+/CD38<sup>-/lo</sup> population of cells probed with MNE6. Fig. 41G shows a table of the detailed analysis. Fig. 41D shows the FACS scatter plot of the CD34+/CD38<sup>-/lo</sup> population of cells probed with MNC2. Fig. 41H shows a table of the detailed analysis.

**[00303]** **Figure 42A-42H** shows photographs of DU145 prostate cancer cells or T47D breast cancer cells that have been treated with either the Fab of anti-MUC1\* antibody MNC2, MNE6, MNC3 or MNC8. The images show that cancer specific antibodies MNC2 and MNE6 effectively kill prostate and breast cancer cells while the monoclonal antibodies MNC3 and MNC8 do not.

**[00304]** **Figure 43** shows a graph of a PCR experiment comparing expression of a wide range of cleavage enzymes expressed in different cells lines, wherein the values have been normalized to those expressed in breast cancer cell line T47D. Cell lines that are compared are prostate cancer cell line DU145, HCT-MUC1-41TR that is a MUC1 negative colon cancer cell line transfected with a MUC1 whose extracellular domain is truncated after 41 tandem repeat units and that is not cleaved to the MUC1\* form, T47D breast cancer cell line and CD34+ bone marrow cells.

**[00305]** **Figure 43** shows a graph of a PCR experiment in which the expression levels of various cleavage enzymes are measured in DU145 prostate cancer cells, HCT116+MUC1FL, also known as HCT-MUC1-18 a cell line expressing full-length MUC1, T47D breast cancer cells, and CD34+ hematopoietic stem cells of the bone marrow. The fold expression is relative to the expression of each cleavage enzyme in T47D breast cancer cells, set as 1.

**[00306]** **Figure 44** shows the graph of the PCR experiment of Figure 43 but with the Y-axis maximum set to 5.

**[00307]** **Figures 45A-45P** show photographs of a CAR T co-culture assay in which the targeting antibody fragment of the CAR is huMNC2scFv wherein CAR44 has a CD8

transmembrane domain, followed by 41BB-3zeta and CAR50 has a CD4 transmembrane domain, followed by 41BB-3zeta. The target cancer cells are: HCT-FLR which is HCT-116 cells transfected with MUC1\*<sub>45</sub> and HCT-MUC1-41TR, which is a stable single cell clone HCT-116 cell line that expresses MUC1 with an extracellular domain truncated after 41 tandem repeats and that does not get cleaved to the MUC1\* form on its own. The HCT-MUC1-41TR cancer cells were also incubated with conditioned media from cells transfected with MMP9 or ADAM17 before co-culture with the CAR T cells. Conditioned media of the MMP9 or ADAM17 expressing cells were also incubated with APMA which is an activator of those cleavage enzymes. The images shown are an overlay of the 4X bright field image and the fluorescent image of the same showing cancer cells dyed with a red CMTMR lipophilic dye. Figs. 45A, 45E, 45I, 45M show photographs of cells co-cultured with untransduced human T cells. Figs. 45B, 45F, 45J, 45N show photographs of cells co-cultured with human T cells transduced with anti-MUC1\* CAR44 at an MOI of 10. Figs. 45C, 45G, 45K, 45O show photographs of cells co-cultured with human T cells transduced with anti-MUC1\* CAR50 at an MOI of 10. Figs. 45D, 45H, 45L, 45P show photographs of cells co-cultured with human T cells transduced with anti-MUC1\* CAR44 at an MOI of 50, which increases transduction efficiency. Figs. 45B, 45C, 45D show that both CAR44 and CAR50 transduced T cells recognized MUC1\* expressed in these cancer cells, bound to them, induced clustering and killed many cancer cells. Figs. 45F, 45G, 45H show that neither CAR44 nor CAR50 transduced T cells recognize full-length MUC1 expressed in HCT-MUC1-41TR cancer cells. There is no T cell induced clustering and the number of cancer cells has not decreased. Figs. 45J, 45K, 45L show that activated MMP9 has cleaved full-length MUC1 to a MUC1\* form that is recognized by both CAR44 and CAR50 transduced T cells. There is clearly visible CAR T cell induced clustering and a decrease in the number of cancer cells as they are killed. Figs. 45N, 45O, 45P show that activated ADAM17 has either not cleaved MUC1 or cleaved it at a position not recognized by MNC2. Neither huMNC2-CAR44 nor huMNC2-CAR50 transduced T cells recognized these cancer cells.

**[00308] Figure 46A-46T** shows photographs of a CAR T co-culture assay in which the targeting antibody fragment of the CAR is MNC2 scFv wherein CAR44 has a CD8 transmembrane domain, followed by 41BB-3zeta and CAR50 has a CD4 transmembrane domain, followed by 41BB-3zeta. The target cancer cells are breast cancer T47D cells that were



also incubated with conditioned media from cells transfected with MMP2, MMP9 or ADAM17 before co-culture with the MNC2-CAR T cells. In some cases, the conditioned media of the MMP2 and MMP9 expressing cells were also incubated with APMA, which is an activator of these cleavage enzymes. The images shown are an overlay of the 4X bright field image and the fluorescent image of the same showing cancer cells dyed with a red CMTMR lipophilic dye. As can be seen, the MNC2-CAR T cells only bind to and attack the target cancer cells that express the cleaved form, MUC1\*.

**[00309] Figures 47A-47I** show photographs of cancer cells co-cultured with anti-MUC1\* CAR T cells, wherein some of the cancer cells were pre-incubated with activated MMP9 prior to co-culture with the CAR T cells. The cancer cells shown in Figs. 47A-47C are MUC1 negative colon cancer cell line HCT-116 that have been stably transfected to express MUC1\*. The cancer cells shown in Figs. 47D-47F are MUC1 positive breast cancer cell line T47Ds that express high levels of both MUC1 full-length and MUC1\*. The cancer cells shown in Figs. 47G-47I are MUC1 positive breast cancer cell line T47Ds that were pre-incubated with activated MMP9. The cells shown in Figs. 47A, 47D and 47G were co-cultured with untransduced human T cells and are the controls. The cells shown in Figs. 47B, 47E and 47H were co-cultured with human T cells that were transduced with huMNC2-CAR44 at an MOI of 10, wherein MOI stands for multiplicity of infection and the higher the MOI the more CARs are expressed on the T cells. The cells shown in Figs. 47C, 47F and 47I were co-cultured with human T cells that were transduced with huMNC2-CAR44 at an MOI of 50. As can be seen in the photographs, the CAR44 T cells bind to the target MUC1\* positive cancer cells, surrounding and killing them. Comparing the photograph Fig. 47I with the others, it can be seen that the cells that were pre-incubated with MMP9 become much more susceptible to CAR T killing when the antibody targeting head of the CAR recognizes MUC1\*. It also demonstrates that MUC1 cleaved by MMP9 is recognized by huMNC2scFv.

**[00310] Figure 48** shows an xCelligence graph of T47D breast cancer cells in co-culture with either untransduced T cells, as a control, or huMNC2-CAR44 T cells over a 45 hour period. After 18 hours of cancer cell growth, a catalytic sub-unit MMP9 was added to some of the cells. At 25 hours, T cells were added. As can be seen, huMNC2-CAR44 T cell killing is greatly improved when the T47D cells are pre-incubated with cleavage enzyme MMP9. In the xCelligence system, target cancer cells, which are adherent, are plated onto electrode array

plates. Adherent cells insulate the electrode and increase the impedance. The number of adherent cancer cells is directly proportional to impedance. T cells are not adherent and do not contribute to impedance. Therefore, increasing impedance reflects growth of cancer cells and decreasing impedance reflects killing of cancer cells.

**[00311]** **Figure 49** shows an xCelligence graph of DU145 prostate cancer cells in co-culture with either untransduced T cells, as a control, or huMNC2-CAR44 T cells over a 45 hour period. After 18 hours of cancer cell growth, a catalytic sub-unit MMP9 was added to some of the cells. At 25 hours, T cells were added. As can be seen, huMNC2-CAR44 T cell killing is not affected by pre-incubation with cleavage enzyme MMP9. DU145 cancer cells express a significantly lower amount of MUC1 which includes the full-length form as well as MUC1\*. The lower density of MUC1 full-length does not sterically hinder T cell access to the membrane proximal MUC1\*.

**[00312]** **Figure 50** shows a bar graph of a PCR experiment measuring the amount of MUC1 expressed by a panel of cell lines and primary cells, comprised of normal cells as well as cancer cells.

**[00313]** **Figure 51A-51B** shows a bar graph of an ELISA assay measuring the amount of interferon gamma, IFN- $\gamma$ , secreted by huMNC2-CAR44 human T cells after co-culture with the normal cells or the HCT-MUC1\* cancer cells for 72 hours. Fig. 51A shows the results of the experiment where the CAR44 T cell to target cell ratio was 1:1. Fig. 51B shows the results of the experiment where the CAR44 T cell to target cell ratio was 0.5:1.

**[00314]** **Figure 52A-52B** shows a bar graph of an ELISA assay measuring the amount of interleukin-2, IL-2, secreted by huMNC2-CAR44 human T cells after co-culture with the normal cells or the HCT-MUC1\* cancer cells for 72 hours. Fig. 52A shows the results of the experiment where the CAR44 T cell to target cell ratio was 1:1. Fig. 52B shows the results of the experiment where the CAR44 T cell to target cell ratio was 0.5:1.

**[00315]** **Figure 53A-53J** shows bar graphs of FACS analysis of live versus dead markers and photographs of normal cells versus cancer cells after co-culture with huMNC2-CAR44 T cells. Fig. 53A.1 shows the bar graph of FACS analysis of live versus dead cells after HCT-MUC1\* cancer cells were co-cultured with huMNC2-CAR44 T cells. Fig. 53A.2 and Fig. 53A.3 show the photographs of the experiment described in Fig. 53A.1. Fig. 53B.1 shows the bar graph of FACS analysis of live versus dead cells after MCF-12A normal breast cells were co-cultured with

huMNC2-CAR44 T cells. Fig. 53B.2 and Fig. 53B.3 show the photographs of the experiment described in Fig. 53B.1. Fig. 53C.1 shows the bar graph of FACS analysis of live versus dead cells after THLE-3 normal liver cells were co-cultured with huMNC2-CAR44 T cells. Fig. 53C.2 and Fig. 53C.3 show the photographs of the experiment described in Fig. 53C.1. Fig. 53D.1 shows the bar graph of FACS analysis of live versus dead cells after T/G HA-HSMC normal heart cells were co-cultured with huMNC2-CAR44 T cells. Fig. 53D.2 and Fig. 53D.3 show the photographs of the experiment described in Fig. 53D.1. Fig. 53E.1 shows the bar graph of FACS analysis of live versus dead cells after Hs1.Tes normal testes cells were co-cultured with huMNC2-CAR44 T cells. Fig. 53E.2 and Fig. 53E.3 show the photographs of the experiment described in Fig. 53E.1. Fig. 53F.1 shows the bar graph of FACS analysis of live versus dead cells after HEK-293 MUC1 negative cells were co-cultured with huMNC2-CAR44 T cells. Fig. 53F.2 and Fig. 53F.3 show the photographs of the experiment described in Fig. 53F.1. Fig. 53G.1 shows the bar graph of FACS analysis of live versus dead cells after HRCE normal kidney cells were co-cultured with huMNC2-CAR44 T cells. Fig. 53G.2 and Fig. 53G.3 show the photographs of the experiment described in Fig. 53G.1. Fig. 53H.1 shows the bar graph of FACS analysis of live versus dead cells after CCD-18Lu normal lung cells were co-cultured with huMNC2-CAR44 T cells. Fig. 53H.2 and Fig. 53H.3 show the photographs of the experiment described in Fig. 53H.1. Fig. 53I.1 shows the bar graph of FACS analysis of live versus dead cells after HBEC-5i normal brain cells were co-cultured with huMNC2-CAR44 T cells. Fig. 53I.2 and Fig. 53I.3 show the photographs of the experiment described in Fig. 53I.1. Fig. 53J.1 shows the bar graph of FACS analysis of live versus dead cells after Hs.738.St/Int normal stomach and intestine cells were co-cultured with huMNC2-CAR44 T cells. Fig. 53J.2 and Fig. 53J.3 show the photographs of the experiment described in Fig. 53J.1.

**[00316] Figure 54** shows photographs of a breast cancer tissue array (CB – insert array number) in which for each patient there is a specimen from the primary tumor plus a specimen from that patient's metastasis. As can be seen in the figure, most often the metastasis expresses more MUC1\* than the primary tumor.

**[00317] Figures 55A-55H** show the cytotoxic effect of huMNC2-CAR44 T cells on MUC1\* positive DU145 prostate cancer cells as measured by a variety of assays. Fig. 55A is a fluorescent photograph of untransduced T cells co-cultured with the prostate cancer cells, wherein granzyme B is stained with a red fluorophore. Fig. 55B shows merging of DAPI and

granzyme B. Fig. 55C is a fluorescent photograph of huMNC2-CAR44 T cells co-cultured with the prostate cancer cells, wherein granzyme B is stained with a red fluorophore. **Fig. 55D** shows merging of DAPI and granzyme B. Fig. 55E is a FACS scan for fluorescently labeled granzyme B for untransduced T cells incubated with the cancer cells. Fig. 55F is a FACS scan showing a positive increase in fluorescently labeled granzyme B for huMNC2-CAR44 T cells incubated with the cancer cells. Fig. 55G is a graph of the mean fluorescent intensity. Fig. 55H is an xCELLigence scan tracking the real-time killing of DU145 cancer cells by huMNC2-CAR44 T cells (blue trace) but not by untransduced T cells (green).

**[00318] Figures 56A-56H** show the cytotoxic effect of huMNC2-CAR44 T cells on MUC1\* positive CAPAN-2 pancreatic cancer cells as measured by a variety of assays. Fig. 56A is a fluorescent photograph of untransduced T cells co-cultured with the pancreatic cancer cells, wherein granzyme B is stained with a red fluorophore. Fig. 56B shows merging of DAPI and granzyme B. Fig. 56C is a fluorescent photograph of huMNC2-CAR44 T cells co-cultured with the pancreatic cancer cells, wherein granzyme B is stained with a red fluorophore. Fig. 56D shows merging of DAPI and granzyme B. Fig. 56E is a FACS scan for fluorescently labeled granzyme B for untransduced T cells incubated with the cancer cells. Fig. 56F is a FACS scan showing a positive increase in fluorescently labeled granzyme B for huMNC2-CAR44 T cells incubated with the cancer cells. Fig. 56G is a graph of the mean fluorescent intensity. Fig. 56H is an xCELLigence scan tracking the real-time killing of CAPAN-2 cancer cells by huMNC2-CAR44 T cells (blue trace) but not by untransduced T cells (green).

**[00319] Figures 57A-57C** show xCELLigence scans tracking the real-time killing of MUC1\* positive cancer cells, but not MUC1\* negative cells, by huMNC2-CAR44 T cells. Fig. 57A shows that huMNC2-CAR44 T cells effectively kill HCT colon cancer cells that have been stably transfected with MUC1\*. Fig. 57B shows that huMNC2-CAR44 T cells have almost no effect on HCT-MUC1-41TR, which is a MUC1 negative cancer cell that has been stably transfected with a MUC1 full-length. In this cell line only about 10% of the cells have MUC1 cleaved to MUC1\*. Fig. 57C shows that huMNC2-CAR44 T cells have no effect on HCT-116 cells, which is a MUC1 negative colon cancer cell line.

**[00320] Figure 58A- 58F** shows photographs NOD/SCID/GAMMA mice in an IVIS instrument measuring photon emission from tumor cells after mice were treated with nothing, PBS, untransduced human T cells or huMNC2-CAR44 T cells. Mice had been injected sub-

cutaneously with HCT-MUC1\* tumor cells that had been made Luciferase positive. Ten (10) minutes before the IVIS photographs were taken, the mice were injected into the intraperitoneal (ip) space with the Luciferase substrate, Luciferin. Fig. 58A shows the tumor bearing mice that had only been treated with phosphate buffered saline, PBS. Fig. 58B shows the tumor bearing mice that had only been treated with untransduced T cells. Fig. 58C shows the tumor bearing mice that had been treated with a single dose of huMNC2-CAR44 T cells. Fig. 58D shows color scale of the images. Fig. 58E shows Kaplan-Meier survival curves of the experiment. Fig. 58F shows a table detailing the molecular makeup of the human T cells that were isolated from the mouse blood after sacrifice.

**[00321] Figure 59A- 59C** shows photographs NOD/SCID/GAMMA mice in an IVIS instrument measuring photon emission from tumor cells after mice were treated with nothing, PBS or huMNC2-CAR44 T cells. Mice had been injected sub-cutaneously with T47D-wt breast cancer cells or T47D+more MUC1\*, which is a mixed population of cells wherein 95% of the cells were T47D cells that had been stably transfected with even more MUC1\*. Both T47D-wt and T47D plus more MUC1\* cells had been made Luciferase positive. Ten (10) minutes before the IVIS photographs were taken, the mice were injected into the intraperitoneal (ip) space with the Luciferase substrate, Luciferin. Fig. 59A shows the tumor bearing mice that had only been treated with phosphate buffered saline, PBS. Fig. 59B shows the T47D-wt tumor bearing mice that had been treated with two (2) doses of huMNC2-CAR44 T cells. Fig. T90.1C shows the T47D-MUC1\* tumor bearing mice that had been treated with two (2) doses of huMNC2-CAR44 T cells.

**[00322] Figure 60A- 60C** shows photographs NOD/SCID/GAMMA mice in an IVIS instrument measuring photon emission from tumor cells after mice were treated with nothing, PBS, untransduced T cells or huMNC2-CAR44 T cells. Mice had been injected sub-cutaneously with a mixed population of 70% T47D-wt breast cancer cells and 30% T47D cells that had been transfected with even more MUC1\*. Both cell types had been made Luciferase positive. Ten (10) minutes before the IVIS photographs were taken, the mice were injected into the intraperitoneal (ip) space with the Luciferase substrate, Luciferin. Fig. 60A shows the tumor bearing mice that had only been treated with phosphate buffered saline, PBS. Fig. 60B shows tumor bearing mice that had only been treated with untransduced T cells. Fig. 60C shows the tumor bearing mice that had been treated with two (2) doses of huMNC2-CAR44 T cells.

**[00323] Figures 61A-61J** show fluorescent photographs of mice taken on an IVIS instrument. NSG (NOD/SCID/GAMMA) immune compromised mice that on Day 0 were sub-cutaneously injected into the flank with 500K human BT-20 cells which are a MUC1\* positive triple negative breast cancer cell line. The cancer cells had been stably transfected with Luciferase. Tumors were allowed to engraft. On Day 6 after IVIS measurement, animals were given a one-time injection of 10 million of either human T cells transduced with huMNC2-scFv-CAR44 or untransduced T cells. 5 million T cells were injected intra-tumor and 5 million were injected into the tail vein. 10 minutes prior to IVIS photographs, mice were IP injected with Luciferin, which fluoresces after cleavage by Luciferase, thus making tumor cells fluoresce. Figs. 61A, 61D, 61G show photographs of mice that were treated with huMNC2-scFv-CAR44 T cells that had been pre-stimulated by co-culturing for 24 hours with 4µm beads to which was attached a synthetic MUC1\*, PSMGFR peptide 24 hours prior to administration: Protocol 1. Figs. 61B, 61E, 61H show photographs of mice that were treated with huMNC2-scFv-CAR44 T cells that had been pre-stimulated by twice co-culturing for 24 hours with MUC1\* positive cancer cells 24 hours prior to administration: Protocol 2. Figs. 61C, 61F, 61I show photographs of mice that were treated with untransduced human T cells. Fig. 61J is a color scale relating fluorescence in photons/second to color.

**[00324] Figures 62A-62M** show fluorescent photographs of mice taken on an IVIS instrument. NSG (NOD/SCID/GAMMA) immune compromised mice that on Day 0 were injected into the intraperitoneal cavity (IP) with 500K human SKOV-3 cells which are a MUC1\* positive ovarian cancer cell line. The cancer cells had been stably transfected with Luciferase. Tumors were allowed to engraft. On Day 4, animals were injected into the intraperitoneal space with 10M either human T cells transduced with huMNC2-scFv-CAR44, untransduced T cells or PBS. On Day 11, animals were injected again except that half the cells were injected into the tail vein and the other half was IP injected. Animals were imaged by IVIS on Days 3, 7, 10 and 15. 10 minutes prior to IVIS photographs, mice were IP injected with Luciferin, which fluoresces after cleavage by Luciferase, thus making tumor cells fluoresce. Figs. 62A, 62D, 62G, and 62J show photographs of mice that were treated with huMNC2-scFv-CAR44 T cells that had been pre-stimulated by co-culturing for 24 hours with 1µm beads to which was attached a synthetic MUC1\*, PSMGFR peptide 24 hours prior to administration. Figs. 62B, 62E, 62H, and 62K show photographs of mice that were treated with untransduced human T cells. Figs. 62C, 62F, 62I, and

62L show photographs of mice that were treated with PBS. Figs. 62A, 62B and 62C are IVIS images taken Day 3 prior to CAR T, T cell or PBS administration. Figs. 62D, 62E and 62F show IVIS images of animals on Day 7, just four (4) days after treatment. Figs. 62G, 62H, and 62I show IVIS images of animals on Day 10. Figs. 62J, 62K, and 62L show IVIS images of animals on Day 15. Fig. 62M is the IVIS color scale relating fluorescence in photons/second to color.

**[00325] Figure 63** shows a graph of an ELISA binding assay in which various monoclonal antibodies are tested for their ability to bind to the PSMGFR peptide, the N-10, C-10, N+20/C-27, or the N+9/C-9 peptide, wherein the concentration of the antibody was at 10ug/mL or 1ug/mL. Note that anti-MUC1\* monoclonal antibodies C2 and E6, which have been demonstrated to be cancer specific, bind to the PSMGFR peptide, still bind if the 10 N-terminal amino acids are missing, but do not bind if the 10 or 9 C-terminal amino acids are missing.

**[00326] Figure 64A-64B** shows a graph of an ELISA binding assay. The antibodies being tested were derived from animals immunized with the PSMGFR peptide. The first selection criteria was to confirm that the antibodies bound to the immunizing PSMGFR peptide. Fig. 64A shows a graph of an ELISA of selected antibodies that were further tested to determine their ability to bind to the PSMGFR peptide, the N-10, the C-10, N+20/C-27, or N+9/C-9 peptide. All the antibodies except 18B4 were able to bind to the N-10 peptide. 18B4 recognized N+20/C-27 but not the N-10 peptide, implying that its cognate epitope lies within the GTINVHDTVET sequence. All except 20A10 and C2 showed some binding to the C-10 and N+9/C-9 peptide, showing that both 20A10 and C2 require the 10 membrane proximal amino acids for binding. C2, which requires the 10 membrane proximal amino acids for binding has been demonstrated to be cancer specific. Fig. 64B shows the sequences of the various peptides. The color of the bars for each antibody in the ELISA graph are color coded to match the deductive cognate sequence, or a portion thereof, of that antibody.

**[00327] Figure 65A-65B** shows a graph of an ELISA binding assay in which various monoclonal antibodies are tested for their ability to bind to the PSMGFR peptide, the N-10, the C-10, N+20/C-27, or N+9/C-9 peptide. The antibodies being tested were derived from animals immunized with the N+20/C-27 peptide. The first selection criteria was to confirm that the antibodies bound to the immunizing N+20/C-27 peptide. Fig. 65A shows a graph of ELISA binding assay that tests the ability of each antibody to bind to various peptides. Although these antibodies were raised against the N+20/C-27 peptide, all but one, 45C11, still bind to the

PSMGFR peptide. The binding of 45C11 is weak but deductive reasoning shows that the cognate epitope must lie within the SNIKFRPGSVV sequence. 1E4 was able to bind to the N+20/C-27 peptide, the PSMGFR and the N-10 peptide, consistent with the idea that its epitope must lie within the QFNQYKTE sequence. Fig. 65B shows the sequences of the various peptides. The color of the bars for each antibody in the ELISA graph are color coded to match the deductive cognate sequence, or a portion thereof, of that antibody.

**[00328]** **Figure 66A-66B** shows a graph of an ELISA binding assay in which various monoclonal antibodies are tested for their ability to bind to the PSMGFR peptide, the N-10, the C-10, N+20/C-27, or N+9/C-9 peptide. The antibodies being tested were derived from animals immunized with the N+9/C-9 peptide. The first selection criteria was to confirm that the antibodies bound to the immunizing N+9/C-9 peptide. Fig. 66A shows a graph of the ELISA assay. All but one, 39H5, were only able to bind to the immunizing peptide, N+9/C-9. 39H5 showed very weak binding to the PSMGFR and N-10 peptide, consistent with the idea that at least a portion of its cognate epitope must lie within the QFNQYKTE sequence. Fig. 66B shows the sequences of the various peptides. The color of the bars for each antibody in the ELISA graph are color coded to match the deductive cognate sequence, or a portion thereof, of that antibody.

**[00329]** **Figure 67A-67D** shows results of ELISA assays to further define antibody epitopes within the MUC1 or MUC1\* extra cellular domain. The antibodies shown in this figure were all generated by immunizing animals with the PSMGFR peptide. Binding assays tested antibodies for their ability to bind to peptides N-19, N-26, N-30, N-10/C-5, N-19/C-5, PSMGFR, N-10 and C-10, which are all subsets of the PSMGFR peptide and numbering refers back to the PSMGFR peptide. Fig. 67A shows the binding of the various antibodies to the various peptides. Fig. 67B shows the sequence of the PSMGFR peptide that has been extended 20 amino acids at the N-terminus. Fig. 67C shows the sequences of the PSMGFR-derived subset peptides. Fig. 67D shows the sequences that comprise all or part of the epitope that is essential for antibody recognition.

**[00330]** **Figure 68A-68D** shows results of ELISA assays to further define antibody epitopes within the MUC1 or MUC1\* extra cellular domain. The antibodies shown in this figure were all generated by immunizing animals with the N+20/C-27 peptide. Binding assays tested antibodies for their ability to bind to peptides N-19, N-26, N-30, N-10/C-5, N-19/C-5, PSMGFR, N-10 and



C-10, which are all subsets of the PSMGFR peptide and numbering refers back to the PSMGFR peptide. Fig. 68A shows the binding of the various antibodies to the various peptides. Fig. 68B shows the sequence of the PSMGFR peptide that has been extended 20 amino acids at the N-terminus. Fig. 68C shows the sequences of the PSMGFR-derived subset peptides. Fig. 68D shows the sequences that comprise all or part of the epitope that is essential for antibody recognition.

**[00331]** **Figures 69A-69D** show results of ELISA assays to further define antibody epitopes within the MUC1 or MUC1\* extra cellular domain. The antibodies shown in this figure were all generated by immunizing animals with the N+9/C-9 peptide. Binding assays tested antibodies for their ability to bind to peptides N-19, N-26, N-30, N-10/C-5, N-19/C-5, PSMGFR, N-10 and C-10, which are all subsets of the PSMGFR peptide and numbering refers back to the PSMGFR peptide. Fig. 69A shows the binding of the various antibodies to the various peptides. Fig. 69B shows the sequence of the PSMGFR peptide that has been extended 20 amino acids at the N-terminus. Fig. 69C shows the sequences of the PSMGFR-derived subset peptides. Fig. 69D shows the sequences that comprise all or part of the epitope that is essential for antibody recognition.

**[00332]** **Figure 70** shows a graph of an ELISA displacement assay. In this experiment, a multi-well plate was coated with the PSMGFR peptide. Recombinant NME7<sub>AB</sub> was allowed to bind to the surface-immobilized PSMGFR peptide. Various antibodies were added, followed by a wash step. The amount of NME7<sub>AB</sub> that remained attached to the PSMGFR coated plate, after antibody competition, was measured by detecting a tag on the NME7<sub>AB</sub>. As a control, anti-NME7<sub>AB</sub> antibodies were also tested for their ability to displace NME7<sub>AB</sub> from the PSMGFR.

**[00333]** **Figure 71A-71H** shows photographs of Western blots in which antibodies are tested for their ability to bind to a linear epitope in full-length MUC1 or MUC1\*. Fig. 71A-71D shows testing of antibodies for ability to bind to a MUC1 negative cell line, HCT-116, or engineered cell lines HCT-MUC1-18, which is a cleavage resistant clone that expresses full-length MUC1, or HCT-MUC1\*, which is engineered to express only the PSMGFR sequence in its extra cellular domain. Fig. 71E-71H shows testing of antibodies for ability to bind to breast cancer cell lines T47D or 1500 aka ZR-75-1. Fig. 71A and Fig. 71E show MNC2, a monoclonal antibody raised against PSMGFR peptide that binds to N-10 but not C-10 variants of the PSMGFR peptide. Fig. 71B and Fig. 71F show MNE6, a monoclonal antibody raised against PSMGFR peptide that

binds to N-10 but not C-10 variants of the PSMGFR peptide. Fig. 71C and Fig. 71G show SDIX, a polyclonal antibody raised against PSMGFR peptide and which binds to the PSMGFR peptide. Fig. 71D and Fig. 71H show VU4H5, a commercially available monoclonal antibody that binds to the tandem repeats of full-length MUC1. As can be seen, neither MNC2 nor MNE6 bind linear epitopes of MUC1 species.

**[00334] Figure 72A-72P** shows photographs of Western blots in which antibodies are tested for their ability to bind to a linear epitope in full-length MUC1 or MUC1\*. All these antibodies were raised against the PSMGFR peptide and bind to the PSMGFR peptide. Fig. 72A-72H shows testing of antibodies for ability to bind to a MUC1 negative cell line, HCT-116, or engineered cell lines HCT-MUC1-18, which is a cleavage resistant clone that expresses full-length MUC1, or HCT-MUC1\*, which is engineered to express only the PSMGFR sequence in its extra cellular domain. Fig. 72I-72P shows testing of antibodies for ability to bind to breast cancer cell lines T47D or 1500 aka ZR-75-1. Fig. 72A and Fig. 72I show 20A10. Fig. 72B and Fig. 72J show 25E6. Fig. 72C and Fig. 72K show 18B4. Fig. 72D and Fig. 72L show 18G12. Fig. 72E and Fig. 72M show 28F9. Fig. 72F and Fig. 72N show 3C2B1. Fig. 72G and Fig. 72O show 5C6F3. Fig. 72H and Fig. 72P show 5C6F3 wherein the blot has been exposed for a longer time period to render more visible the MUC1\* specific bands. As can be seen, antibodies 25E6, 18B4 and to a degree 5C6F3 recognize linear epitopes but 20A10, 3C2B1, 18G12 and 28F9 do not.

**[00335] Figure 73A-73J** shows photographs of Western blots in which antibodies are tested for their ability to bind to a linear epitope in full-length MUC1 or MUC1\*. All these antibodies were raised against the N+20/C-27 variant of the PSMGFR peptide and bind to the N+20/C-27 peptide. Fig. 73A-73E shows testing of antibodies for ability to bind to a MUC1 negative cell line, HCT-116, or engineered cell lines HCT-MUC1-18, which is a cleavage resistant clone that expresses full-length MUC1, or HCT-MUC1\*, which is engineered to express only the PSMGFR sequence in its extra cellular domain. Fig. 73F-73J shows testing of antibodies for ability to bind to breast cancer cell lines T47D or 1500 aka ZR-75-1. Fig. 73A and Fig. 73F show 1E4. Fig. 73B and Fig. 73G show 45C11. Fig. 73C and Fig. 73H show 31A1. Fig. 73D and Fig. 73I show 32C1. Fig. 73E and Fig. 73J show 29H1. As can be seen, antibodies 31A1 and 32C1 recognize linear epitopes.

**[00336] Figure 74A-74H** shows photographs of Western blots in which antibodies are tested for their ability to bind to a linear epitope in full-length MUC1 or MUC1\*. All these antibodies

were raised against the N+9/C-9 variant of the PSMGFR peptide and bind to the N+9/C-9 peptide. Fig. 74A-74D shows testing of antibodies for ability to bind to a MUC1 negative cell line, HCT-116, or engineered cell lines HCT-MUC1-18, which is a cleavage resistant clone that expresses full-length MUC1, or HCT-MUC1\*, which is engineered to express only the PSMGFR sequence in its extra cellular domain. Fig. 74E-74H shows testing of antibodies for ability to bind to breast cancer cell lines T47D or 1500 aka ZR-75-1. Fig. 74A and Fig. 74E show 8A9. Fig. 74B and Fig. 74F show 17H6. Fig. 74C and Fig. 74G show 3C5. Fig. 74D and Fig. 74H show 39H5.

**[00337] Figure 75A-75P** show graphs of FACS analysis. HCT-MUC1-18 cells, which express full-length MUC1, were incubated with a catalytically active MMP9 or MMP2 for 24 hours, incubated with an antibody of the invention and then analyzed by FACS to see if the antibody bound to the MMP9 or the MMP2 cleaved form of MUC1. Note that the first bar of each graph shows that none of the antibodies binds to full-length MUC1 in the absence of cleavage. Each bar graph is labeled with both the name of the antibody used in that assay and its cognate epitope. The order of the graphs from right to left corresponds to the distance the from the cell surface of the antibody's cognate epitope. Fig. 75A shows antibody 1E4. Fig. 75B shows antibody 28F9. Fig. 75C shows antibody 18G12. Fig. 75D shows antibody 25E6. Fig. 75E shows antibody 20A10. Fig. 75F shows antibody 3C5. Fig. 75G shows antibody 29H1. Fig. 75H shows antibody 32C1. Fig. 75I shows antibody 31A1. Fig. 75J shows antibody 18B4. Fig. 75K shows antibody 45C11. Fig. 75L shows antibody 8A9. Fig. 75M shows antibody 17H6. Fig. 75N shows antibody 39H5. Fig. 75O shows antibody 3C2B1. Fig. 75P shows antibody 5C6F3.

**[00338] Figure 76A-76J** show graphs of FACS analyses of reference antibodies MNC2, "C2", and VU4H5 binding to either the MUC1-negative cell line HCT-116, HCTs transfected with MUC1\*, "HCT-MUC1\*", a cleavage resistant single cell clone of HCTs transfected with MUC1 full-length, "HCT-MUC1-18", and MNC2 binding to breast cancer cells line T47D or breast cancer cell line 1500 also known as ZR-75-1. MNC2 binds to an ectopic binding site on the extra cellular domain of MUC1\*, within the membrane proximal portion of the PSMGFR sequence. The MNC2 binding site is only available after cleavage and release of the bulk of the extra cellular domain comprising the tandem repeat domain. VU4H5 binds to hundreds of repeating epitopes in the tandem repeat domain. Fig. 76A-76E show percent binding and Fig. 76F-76J show Mean Fluorescent Intensity or MFI.

**[00339]** **Figure 77A-77N** show graphs of FACS analyses of reference antibody MNC2, “C2”, binding to a panel of cancer cell lines that are MUC1\* positive, with the exception of MDA-MB-231, which expresses MUC1 and MUC1\* at a level that is so low that it is often used as a negative control. MNC2 binds to an ectopic binding site on the extra cellular domain of MUC1\*, within the membrane proximal portion of the PSMGFR sequence. The MNC2 binding site is only available after cleavage and release of the bulk of the extra cellular domain comprising the tandem repeat domain. Fig. 77A-77G show percent binding and Fig. 77H-77N show Mean Fluorescent Intensity or MFI. Fig. 77A and 77H show the antibodies binding to lung cancer cell line NCI-H292. Fig. 77B and 77I show the antibodies binding to lung cancer cell line NCI-H1975. Fig. 77C and 77J show the antibodies binding to ovarian cancer cell line SKOV-3. Fig. 77D and 77K show the antibodies binding to pancreatic cancer cell line HPAF-II. Fig. 77E and 77L show the antibodies binding to pancreatic cancer cell line Capan-1. Fig. 77F and 77M show the antibodies binding to prostate cancer cell line DU145. Fig. 77G and 77N show the antibodies binding to breast cancer cell line MDA-MB-231, which is nearly MUC1 and MUC1\* negative.

**[00340]** **Figure 78A-78C** shows a color coded schematic of the basic PSMGFR sequence that has been extended or deleted at both the N- and C-termini. Antibodies of the invention were tested against this subset of peptides to further refine the epitopes to which each antibody binds or the critical amino acids within the epitope to which each antibody binds. Fig. 78A is an aligned schematic of the various subsets of peptides. Fig. 78B lists the antibodies that bind to each of the color coded sequences. Fig. 78C lists the cancer cell lines that each antibody recognizes.

**[00341]** **Figure 79A-79I** shows color coded graphs that resulted from FACS analyses of each antibody binding to T47D breast cancer cells and their respective cognate sequences within the N-terminally extended PSMGFR sequence. Fig. 79A-79D are FACS graphs showing the percent cells that were recognized by each antibody. Fig. 79E-79H are FACS graphs showing the Mean Fluorescence Intensity, MFI, of each antibody. Fig. 79A and Fig. 79E show the FACS graph of antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 79B and Fig. 79F show the FACS graph of antibodies that were generated by immunizing with the N+20/C-27 peptide. Fig. 79C and Fig. 79G show the FACS graph of antibodies that were generated by immunizing with the N+9/C-9 peptide. Fig. 79D and Fig. 79H also show the FACS graph of

antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 79I shows the PSMGFR sequence that is extended at the N-terminus by 20 amino acids.

**[00342]** **Figure 80A-80I** shows color coded graphs that resulted from FACS analyses of each antibody binding to 1500, also known as ZR-75-1, breast cancer cells and their respective cognate sequences within the N-terminally extended PSMGFR sequence. Fig. 80A-80C are FACS graphs showing the percent cells that were recognized by each antibody. Fig. 80D-80F are FACS graphs showing the Mean Fluorescence Intensity, MFI, of each antibody. Fig. 80A, Fig. 80E Fig. 80D and Fig. 80H show the FACS graph of antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 80B and Fig. 80F show the FACS graph of antibodies that were generated by immunizing with the N+20/C-27 peptide. Fig. 80C and Fig. 80G show the FACS graph of antibodies that were generated by immunizing with the N+9/C-9 peptide. Fig. 80I shows the PSMGFR sequence that is extended at the N-terminus by 20 amino acids.

**[00343]** **Figure 81A-81G** shows color coded graphs that resulted from FACS analyses of each antibody binding to NCI-H292 lung cancer cells and their respective cognate sequences within the N-terminally extended PSMGFR sequence. Fig. 81A-81C are FACS graphs showing the percent cells that were recognized by each antibody. Fig. 81D-81F are FACS graphs showing the Mean Fluorescence Intensity, MFI, of each antibody. Fig. 81A and Fig. 81D show the FACS graph of antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 81B and Fig. 81E show the FACS graph of antibodies that were generated by immunizing with the N+20/C-27 peptide. Fig. 81C and Fig. 81F show the FACS graph of antibodies that were generated by immunizing with the N+9/C-9 peptide. Fig. 81G shows the PSMGFR sequence that is extended at the N-terminus by 20 amino acids.

**[00344]** **Figure 82A-82G** shows color coded graphs that resulted from FACS analyses of each antibody binding to NCI-H1975 lung cancer cells and their respective cognate sequences within the N-terminally extended PSMGFR sequence. Fig. 82A-82C are FACS graphs showing the percent cells that were recognized by each antibody. Fig. 82D-82F are FACS graphs showing the Mean Fluorescence Intensity, MFI, of each antibody. Fig. 82A and Fig. 82D show the FACS graph of antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 82B and Fig. 82E show the FACS graph of antibodies that were generated by immunizing with the N+20/C-27 peptide. Fig. 82C and Fig. 82F show the FACS graph of antibodies that were

generated by immunizing with the N+9/C-9 peptide. Fig. 82G shows the PSMGFR sequence that is extended at the N-terminus by 20 amino acids.

**[00345]** **Figure 83A-83G** shows color coded graphs that resulted from FACS analyses of each antibody binding to SKOV-3 ovarian cancer cells and their respective cognate sequences within the N-terminally extended PSMGFR sequence. Fig. 83A-83C are FACS graphs showing the percent cells that were recognized by each antibody. Fig. 83D-83F are FACS graphs showing the Mean Fluorescence Intensity, MFI, of each antibody. Fig. 83A and Fig. 83D show the FACS graph of antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 83B and Fig. 83E show the FACS graph of antibodies that were generated by immunizing with the N+20/C-27 peptide. Fig. 83C and Fig. 83F show the FACS graph of antibodies that were generated by immunizing with the N+9/C-9 peptide. Fig. 83G shows the PSMGFR sequence that is extended at the N-terminus by 20 amino acids.

**[00346]** **Figure 84A-84G** shows color coded graphs that resulted from FACS analyses of each antibody binding to DU145 prostate cancer cells and their respective cognate sequences within the N-terminally extended PSMGFR sequence. Fig. 84A-84C are FACS graphs showing the percent cells that were recognized by each antibody. Fig. 84D-84F are FACS graphs showing the Mean Fluorescence Intensity, MFI, of each antibody. Fig. 84A and Fig. 84D show the FACS graph of antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 84B and Fig. 84E show the FACS graph of antibodies that were generated by immunizing with the N+20/C-27 peptide. Fig. 84C and Fig. 84F show the FACS graph of antibodies that were generated by immunizing with the N+9/C-9 peptide. Fig. 84G shows the PSMGFR sequence that is extended at the N-terminus by 20 amino acids.

**[00347]** **Figure 85A-85G** shows color coded graphs that resulted from FACS analyses of each antibody binding to HPAF-II pancreatic cancer cells and their respective cognate sequences within the N-terminally extended PSMGFR sequence. Fig. 85A-85C are FACS graphs showing the percent cells that were recognized by each antibody. Fig. 85D-85F are FACS graphs showing the Mean Fluorescence Intensity, MFI, of each antibody. Fig. 85A and Fig. 85D show the FACS graph of antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 85B and Fig. 85E show the FACS graph of antibodies that were generated by immunizing with the N+20/C-27 peptide. Fig. 85C and Fig. 85F show the FACS graph of antibodies that were

generated by immunizing with the N+9/C-9 peptide. Fig. 85G shows the PSMGFR sequence that is extended at the N-terminus by 20 amino acids.

**[00348]** **Figure 86A-86G** shows color coded graphs that resulted from FACS analyses of each antibody binding to Capan-1 pancreatic cancer cells and their respective cognate sequences within the N-terminally extended PSMGFR sequence. Fig. 86A-86C are FACS graphs showing the percent cells that were recognized by each antibody. Fig. 86D-86F are FACS graphs showing the Mean Fluorescence Intensity, MFI, of each antibody. Fig. 86A and Fig. 86D show the FACS graph of antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 86B and Fig. 86E show the FACS graph of antibodies that were generated by immunizing with the N+20/C-27 peptide. Fig. 86C and Fig. 86F show the FACS graph of antibodies that were generated by immunizing with the N+9/C-9 peptide. Fig. 86G shows the PSMGFR sequence that is extended at the N-terminus by 20 amino acids.

**[00349]** **Figure 87A-87G** shows color coded graphs that resulted from FACS analyses of each antibody binding to MDA-MB-231 breast cancer cells, which are nearly MUC1 negative, and their respective cognate sequences within the N-terminally extended PSMGFR sequence. Fig. 87A-87C are FACS graphs showing the percent cells that were recognized by each antibody. Fig. 87D-87F are FACS graphs showing the Mean Fluorescence Intensity, MFI, of each antibody. Fig. 87A and Fig. 87D show the FACS graph of antibodies that were generated by immunizing with the PSMGFR peptide. Fig. 87B and Fig. 87E show the FACS graph of antibodies that were generated by immunizing with the N+20/C-27 peptide. Fig. 87C and Fig. 87F show the FACS graph of antibodies that were generated by immunizing with the N+9/C-9 peptide. Fig. 87G shows the PSMGFR sequence that is extended at the N-terminus by 20 amino acids.

**[00350]** **Figure 88A-88L** show photographs of normal liver tissue specimens, each from the same donor but stained with a different antibody of the invention. Fig. 88A-88F show the entire tissue core. Fig. 88G-88L show the 40X magnification of a particular area of the tissue. The tissues are ordered from right to left with antibodies that bind to the most membrane proximal, that is to say most C-terminal portion of the PSMGFR peptide, on the right and antibodies that bind to the most N-terminal portions of the MUC1 extra cellular domain, even beyond the PSMGFR region, on the left. As can be seen in the figure, the most cancer-specific antibodies are those that bind to the more membrane proximal portions of the PSMGFR sequence and

antibodies that bind to the most distal, N-terminal portions lose cancer specificity, with those antibodies that bind to epitopes outside of the PSMGFR having lost all cancer specificity.

**[00351]** **Figure 89A-89H** show photographs of normal heart tissue specimens, stained with different antibodies of the invention. Fig. 89A-89D show the entire tissue core. Fig. 89E-89H show the 40X magnification of a particular area of the tissue. Fig. 89A and Fig. 89E show staining with MNC2-scFv. Fig. 89B and Fig. 89F show staining with MNE6. Fig. 89C and Fig. 89G show staining with 20A10. Fig. 89D and Fig. 89H show staining with 3C2B1. These antibodies bind to an epitope that comprises all or part of the sequence FPFS or PFPFSAQSGA. All these antibodies are all able to bind to the PSMGFR peptide, bind to the N-10 peptide but do not bind to the C-10 peptide. In addition, these antibodies disrupt the binding of NME7<sub>AB</sub> to the MUC1\* extra cellular domain as exemplified by the PSMGFR peptide. Further, these antibodies recognize a MUC1 cleavage product when the cleavage enzyme is MMP9. As can be seen in the figure, these antibodies show no binding to normal heart tissue.

**[00352]** **Figure 90A-90D** show photographs of normal heart tissue specimens, stained with different antibodies of the invention. Fig. 90A-90B show the entire tissue core. Fig. 90C-90D show the 40X magnification of a particular area of the tissue. Fig. 90A and Fig. 90C show staining with MNC3. Fig. 90B and Fig. 90D show staining with 25E6. These antibodies bind to an epitope that comprises all or part of the sequence ASRYNLT. These antibodies are all able to bind to the PSMGFR peptide, bind to the N-10 peptide but also bind to the C-10 peptide.

**[00353]** **Figure 91A-91B** show photographs of normal heart tissue specimens, stained with an antibody of the invention 1E4. Fig. 91A show the entire tissue core. Fig. 91B show the 40X magnification of a particular area of the tissue. Antibody 1E4 binds to an epitope that comprises all or part of the sequence QFNQYKTEA. Antibody 1E4 can bind to the N-10 peptide but also binds to the C-10 peptide. As can be seen in the figure, 1E4 binds to normal heart tissue.

**[00354]** **Figure 92A-92H** show photographs of normal heart tissue specimens, stained with different antibodies of the invention. Fig. 92A-92D show the entire tissue core. Fig. 92E-92H show the 40X magnification of a particular area of the tissue. Fig. 92A and Fig. 92E show staining with 18B4. Fig. 92B and Fig. 92F show staining with 31A1. Fig. 92C and Fig. 92G show staining with 32C1. Fig. 92D and Fig. 92H show staining with 29H1. These antibodies bind to an epitope that comprises all or part of the sequence GTINVHDTVET, which is the most N-terminal part of the PSMGFR peptide. None of these antibodies are able to bind to the N-10



peptide. As can be seen in the figure, all of these antibodies except 18B4 show bind to normal heart tissue.

**[00355]** **Figure 93A-93D** show photographs of normal heart tissue specimens, stained with antibodies of the invention. Fig. 93A-93B show the entire tissue core. Fig. 93C-93D show the 40X magnification of a particular area of the tissue. Fig. 93A and Fig. 93C show staining with antibody 8A9. Fig. 93B and Fig 93D show staining with antibody 17H6. Both antibodies bind to an epitope that that is outside of the PSMGFR region and comprises all or part of the sequence VQLTLAFRE. As can be seen in the figure, both antibodies show strong binding to normal heart tissue.

**[00356]** **Figure 94A-94B** show photographs of normal heart tissue specimens, stained with an antibody of the invention 45C11. Fig. 94A show the entire tissue core. Fig. 94B show the 40X magnification of a particular area of the tissue. Antibody 45C11 binds to an epitope that is outside of the PSMGFR region and comprises all or part of the sequence **SNIKFRPGSVV**. Antibody 45C11 cannot bind to the N-10 peptide. As can be seen in the figure, 45C11 binds strongly to normal heart tissue.

**[00357]** **Figure 95A-95H** show photographs of normal liver tissue specimens, stained with different antibodies of the invention. Fig. 95A-95D show the entire tissue core. Fig. 95E-95HL show the 40X magnification of a particular area of the tissue. Fig. 95A and Fig. 95E show staining with MNC2-scFv. Fig. 95B and Fig. 95F show staining with MNE6. Fig. 95C and Fig. 95G show staining with 20A10. Fig. 95D and Fig. 95H show staining with 3C2B1. These antibodies bind to an epitope that comprises all or part of the sequence FPFS or PFPFSAQSGA. All these antibodies are all able to bind to the PSMGFR peptide, bind to the N-10 peptide but do not bind to the C-10 peptide. In addition, these antibodies disrupt the binding of NME7<sub>AB</sub> to the MUC1\* extra cellular domain as exemplified by the PSMGFR peptide. Further, these antibodies recognize a MUC1 cleavage product when the cleavage enzyme is MMP9. As can be seen in the figure, these antibodies show no binding to normal liver tissue.

**[00358]** **Figure 96A-96D** show photographs of normal liver tissue specimens, stained with different antibodies of the invention. Fig. 96A-96B show the entire tissue core. Fig. 96C-96D show the 40X magnification of a particular area of the tissue. Fig. 96A and Fig. 96C show staining with MNC3. Fig. 96B and Fig. 96D show staining with 25E6. These antibodies bind to

an epitope that comprises all or part of the sequence ASRYNLT. These antibodies are all able to bind to the PSMGFR peptide, bind to the N-10 peptide but also bind to the C-10 peptide.

**[00359]** **Figure 97A-97B** show photographs of normal liver tissue specimens, stained with an antibody of the invention 1E4. Fig. 97A show the entire tissue core. Fig. 97B show the 40X magnification of a particular area of the tissue. Antibody 1E4 binds to an epitope that comprises all or part of the sequence QFNQYKTEA. Antibody 1E4 can bind to the N-10 peptide but also binds to the C-10 peptide. As can be seen in the figure, 1E4 binds to normal liver tissue.

**[00360]** **Figure 98A-98H** show photographs of normal liver tissue specimens, stained with different antibodies of the invention. Fig. 98A-98D show the entire tissue core. Fig. 98E-98H show the 40X magnification of a particular area of the tissue. Fig. 98A and Fig. 98E show staining with 18B4. Fig. 98B and Fig. 98F show staining with 31A1. Fig. 98C and Fig. 98G show staining with 32C1. Fig. 98D and Fig. 98H show staining with 29H1. These antibodies bind to an epitope that comprises all or part of the sequence GTINVHDTVET, which is the most N-terminal part of the PSMGFR peptide. None of these antibodies are able to bind to the N-10 peptide. As can be seen in the figure, 32C1 shows some binding to normal liver and 29H1 shows extremely strong binding to normal liver tissue.

**[00361]** **Figure 99A-99D** show photographs of normal liver tissue specimens, stained with antibodies of the invention. Fig. 99A-99B show the entire tissue core. Fig. 99C-99D show the 40X magnification of a particular area of the tissue. Fig. 99A and Fig. 99C show staining with antibody 8A9. Fig. 99B and Fig. 99D show staining with antibody 17H6. Both antibodies bind to an epitope that that is outside of the PSMGFR region and comprises all or part of the sequence VQLTLAFRE. As can be seen in the figure, 8A9 shows strong binding to normal liver tissue. 17H6 is a weak antibody and it is possible that it was not used at a high enough concentration in this study.

**[00362]** **Figure 100A-100B** show photographs of normal liver tissue specimens, stained with an antibody of the invention 45C11. Fig. 100A show the entire tissue core. Fig. 100B show the 40X magnification of a particular area of the tissue. Antibody 45C11 binds to an epitope that is outside of the PSMGFR region and comprises all or part of the sequence **SNIKFRPGSVV**. Antibody 45C11 cannot bind to the N-10 peptide. As can be seen in the figure, 45C11 binds strongly to normal liver tissue.

**[00363]** **Figure 101A-101H** show photographs of normal lung tissue specimens, stained with different antibodies of the invention. Fig. 101A-101D show the entire tissue core. Fig. 101E-101H show the 40X magnification of a particular area of the tissue. Fig. 101A and Fig. 101E show staining with MNC2-scFv. Fig. 101B and Fig. 101F show staining with MNE6. Fig. 101C and Fig. 101G show staining with 20A10. Fig. 101D and Fig. 101H show staining with 3C2B1. These antibodies bind to an epitope that comprises all or part of the sequence FPFS or PFPFSAQSGA. All these antibodies are all able to bind to the PSMGFR peptide, bind to the N-10 peptide but do not bind to the C-10 peptide. In addition, these antibodies disrupt the binding of NME7<sub>AB</sub> to the MUC1\* extra cellular domain as exemplified by the PSMGFR peptide. Further, these antibodies recognize a MUC1 cleavage product when the cleavage enzyme is MMP9. As can be seen in the figure, these antibodies show no binding to normal lung tissue.

**[00364]** **Figure 102A-102D** show photographs of normal lung tissue specimens, stained with different antibodies of the invention. Fig. 102A-102B show the entire tissue core. Fig. 102C-102D show the 40X magnification of a particular area of the tissue. Fig. 102A and Fig. 102C show staining with MNC3. Fig. 102B and Fig. 102D show staining with 25E6. These antibodies bind to an epitope that comprises all or part of the sequence ASRYNLT. These antibodies are all able to bind to the PSMGFR peptide, bind to the N-10 peptide but also bind to the C-10 peptide.

**[00365]** **Figure 103A-103B** show photographs of normal lung tissue specimens, stained with an antibody of the invention 1E4. Fig. 103A show the entire tissue core. Fig. 103B show the 40X magnification of a particular area of the tissue. Antibody 1E4 binds to an epitope that comprises all or part of the sequence QFNQYKTEA. Antibody 1E4 can bind to the N-10 peptide but also binds to the C-10 peptide.

**[00366]** **Figure 104A-104H** show photographs of normal lung tissue specimens, stained with different antibodies of the invention. Fig. 104A-104D show the entire tissue core. Fig. 104E-104H show the 40X magnification of a particular area of the tissue. Fig. 104A and Fig. 104E show staining with 18B4. Fig. 104B and Fig. 104F show staining with 31A1. Fig. 104C and Fig. 104G show staining with 32C1. Fig. 104D and Fig. 104H show staining with 29H1. These antibodies bind to an epitope that comprises all or part of the sequence GTINVHDTVET, which is the most N-terminal part of the PSMGFR peptide. None of these antibodies are able to bind to the N-10 peptide. As can be seen in the figure, all these antibodies show strong binding to normal lung tissue.

**[00367]** **Figure 105A-105D** show photographs of normal lung tissue specimens, stained with antibodies of the invention. Fig. 105A-105B show the entire tissue core. Fig. 105C-105D show the 40X magnification of a particular area of the tissue. Fig. 105A and Fig. 105C show staining with antibody 8A9. Fig. 105B and Fig. 105D show staining with antibody 17H6. Both antibodies bind to an epitope that is outside of the PSMGFR region and comprises all or part of the sequence VQLTLAFRE. As can be seen in the figure, 8A9 shows strong binding to normal lung tissue. 17H6 is a weak antibody and it is possible that it was not used at a high enough concentration in this study.

**[00368]** **Figure 106A-106B** show photographs of normal lung tissue specimens, stained with an antibody of the invention 45C11. Fig. 106A show the entire tissue core. Fig. 106B show the 40X magnification of a particular area of the tissue. Antibody 45C11 binds to an epitope that is outside of the PSMGFR region and comprises all or part of the sequence **SNIKFRPGSVV**. Antibody 45C11 cannot bind to the N-10 peptide. As can be seen in the figure, 45C11 binds to normal lung tissue.

**[00369]** **Figure 107A-107H** show photographs of normal bone marrow tissue specimens, stained with different antibodies of the invention. Fig. 107A-107D show the entire tissue core. Fig. 107E-107H show the 40X magnification of a particular area of the tissue. Fig. 107A and Fig. 107E show staining with MNC2-scFv. Fig. 107B and Fig. 107F show staining with MNE6. Fig. 107C and Fig. 107G show staining with 20A10. Fig. 107D and Fig. 107H show staining with 3C2B1. These antibodies bind to an epitope that comprises all or part of the sequence FPFSS or PFPFSAQSGA. All these antibodies are all able to bind to the PSMGFR peptide, bind to the N-10 peptide but do not bind to the C-10 peptide. In addition, these antibodies disrupt the binding of NME7<sub>AB</sub> to the MUC1\* extra cellular domain as exemplified by the PSMGFR peptide. Further, these antibodies recognize a MUC1 cleavage product when the cleavage enzyme is MMP9. As can be seen in the figure, these antibodies show no binding to normal bone marrow tissue.

**[00370]** **Figure 108A-108D** show photographs of normal bone marrow tissue specimens, stained with different antibodies of the invention. Fig. 108A-108B show the entire tissue core. Fig. 108C-108D show the 40X magnification of a particular area of the tissue. Fig. 108A and Fig. 108C show staining with MNC3. Fig. 108B and Fig. 108D show staining with 25E6. These antibodies bind to an epitope that comprises all or part of the sequence ASRYNLT. These

antibodies are all able to bind to the PSMGFR peptide, bind to the N-10 peptide but also bind to the C-10 peptide.

**[00371]** **Figure 109A-109B** show photographs of normal bone marrow tissue specimens, stained with an antibody of the invention 1E4. Fig. 109A show the entire tissue core. Fig. 109B show the 40X magnification of a particular area of the tissue. Antibody 1E4 binds to an epitope that comprises all or part of the sequence QFNQYKTEA. Antibody 1E4 can bind to the N-10 peptide but also binds to the C-10 peptide. 1E4 binds to normal bone marrow.

**[00372]** **Figure 110A-110H** show photographs of normal bone marrow tissue specimens, stained with different antibodies of the invention. Fig. 110A-110D show the entire tissue core. Fig. 110E-110H show the 40X magnification of a particular area of the tissue. Fig. 110A and Fig. 110E show staining with 18B4. Fig. 110B and Fig. 110F show staining with 31A1. Fig. 110C and Fig. 110G show staining with 32C1. Fig. 110D and Fig. 110H show staining with 29H1. These antibodies bind to an epitope that comprises all or part of the sequence GTINVHDTVET, which is the most N-terminal part of the PSMGFR peptide. None of these antibodies are able to bind to the N-10 peptide. As can be seen in the figure, all these antibodies show strong binding to normal bone marrow tissue.

**[00373]** **Figure 111A-111D** show photographs of normal bone marrow tissue specimens, stained with antibodies of the invention. Fig. 111A-111B show the entire tissue core. Fig. 111C-111D show the 40X magnification of a particular area of the tissue. Fig. 111A and Fig. 111C show staining with antibody 8A9. Fig. 111B and Fig. 111D show staining with antibody 17H6. Both antibodies bind to an epitope that is outside of the PSMGFR region and comprises all or part of the sequence VQLTLAFRE. As can be seen in the figure, 8A9 shows strong binding to normal bone marrow tissue. 17H6 is a weak antibody and it is possible that it was not used at a high enough concentration in this study.

**[00374]** **Figure 112A-112B** show photographs of normal bone marrow tissue specimens, stained with an antibody of the invention 45C11. Fig. 112A show the entire tissue core. Fig. 112B show the 40X magnification of a particular area of the tissue. Antibody 45C11 binds to an epitope that is outside of the PSMGFR region and comprises all or part of the sequence **SNIKFRPGSVV**. Antibody 45C11 cannot bind to the N-10 peptide. As can be seen in the figure, 45C11 binds to normal bone marrow tissue.

**[00375]** **Figure 113A-113C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 20A10 at 0.25ug/mL. Fig. 113A shows photographs of the tissue micro array. Fig. 113B shows map of the array with abbreviated tissue descriptors. Fig. 113C detailed description of the tissue micro array with non-identifying donor data.

**[00376]** **Figure 114A-114X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 20A10 at 0.25ug/mL, magnified to 6X and 20X. Fig. 114A and Fig. 114E are adrenal gland. Fig. 114B and Fig. 114F are breast. Fig. 114C and Fig. 114G are fallopian tube. Fig. 114D and Fig. 114H are kidney. Fig. 114I and Fig. 114M are heart muscle. Fig. 114J and Fig. 114N are liver. Fig. 114K and Fig. 114O are lung. Fig. 114L and Fig. 114P are ureter. Fig. 114Q and Fig. 114U are eye. Fig. 114R and Fig. 114V are cerebral cortex. Fig. 114S and Fig. 114W are bone marrow. Fig. 114T and Fig. 114X are skeletal muscle.

**[00377]** **Figure 115A-115C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 20A10 at 0.25ug/mL. Fig. 115A shows photographs of the tissue micro array. Fig. 115B shows map of the array with abbreviated tissue descriptors. Fig. 115C detailed description of the tissue micro array with non-identifying donor data.

**[00378]** **Figure 116A-116F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 20A10 at 0.25ug/mL, magnified to 6X and 20X. Fig. 116A and Fig. 116D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 116B and Fig. 116E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 116C and Fig. 116F are photographs of a Grade 2 invasive ductal carcinoma.

**[00379]** **Figure 117A-117C** shows photographs, array map and description of pancreatic cancer tissue array PA805c stained with the anti-PSMGFR antibody 20A10 at 0.25ug/mL. Fig. 117A shows photographs of the tissue micro array. Fig. 117B shows map of the array with abbreviated tissue descriptors. Fig. 117C detailed description of the tissue micro array with non-identifying donor data.

**[00380]** **Figure 118A-118F** shows photographs of specific tissues from pancreatic cancer tissue array PA805c stained with the anti-PSMGFR antibody 20A10 at 0.25ug/mL, magnified to 6X and 20X. Fig. 118A and Fig. 118D are photographs of a Grade 2 papillary adenocarcinoma.

Fig. 118B and Fig. 118E are photographs of a Grade 2-3 ductal carcinoma. Fig. 118C and Fig. 118F are photographs of a Grade 3 invasive adenocarcinoma.

**[00381]** **Figure 119A-119C** shows photographs, array map and description of esophageal cancer tissue array BC001113 stained with the anti-PSMGFR antibody 20A10 at 0.25ug/mL. Fig. 119A shows photographs of the tissue micro array. Fig. 119B shows map of the array with abbreviated tissue descriptors. Fig. 119C detailed description of the tissue micro array with non-identifying donor data.

**[00382]** **Figure 120A-120F** shows photographs of specific tissues from esophageal cancer tissue array BC001113 stained with the anti-PSMGFR antibody 20A10 at 0.25ug/mL, magnified to 6X and 20X. Fig. 120A and Fig. 120D are photographs of the specimen at position A1. Fig. 120B and Fig. 120E are photographs of the specimen at position A7. Fig. 120C and Fig. 120F are photographs of the specimen at position A8.

**[00383]** **Figure 121A-121C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL. Fig. 121A shows photographs of the tissue micro array. Fig. 121B shows map of the array with abbreviated tissue descriptors. Fig. 121C detailed description of the tissue micro array with non-identifying donor data.

**[00384]** **Figure 122A-122X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL, magnified to 6X and 20X. Fig. 122A and Fig. 122E are adrenal gland. Fig. 122B and Fig. 122F are breast. Fig. 122C and Fig. 122G are fallopian tube. Fig. 122D and Fig. 122H are kidney. Fig. 122I and Fig. 122M are heart muscle. Fig. 122J and Fig. 122N are liver. Fig. 122K and Fig. 122O are lung. Fig. 122L and Fig. 122P are ureter. Fig. 122Q and Fig. 122U are eye. Fig. 122R and Fig. 122V are cerebral cortex. Fig. 122S and Fig. 122W are bone marrow. Fig. 122T and Fig. 122X are skeletal muscle.

**[00385]** **Figure 123A-123C** shows photographs, array map and description of pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL. Fig. 123A shows photographs of the tissue micro array. Fig. 123B shows map of the array with abbreviated tissue descriptors. Fig. 123C detailed description of the tissue micro array with non-identifying donor data.

[00386] **Figure 124A-124F** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL, magnified to 6X and 20X. Fig. 124A and Fig. 124D are photographs of a Grade 2 adenocarcinoma. Fig. 124B and Fig. 124E are photographs of a Grade 2 adenocarcinoma. Fig. 124C and Fig. 124F are photographs of a Grade 2 adenocarcinoma.

[00387] **Figure 125A-125C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL. Fig. 125A shows photographs of the tissue micro array. Fig. 125B shows map of the array with abbreviated tissue descriptors. Fig. 125C detailed description of the tissue micro array with non-identifying donor data.

[00388] **Figure 126A-126F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL, magnified to 6X and 20X. Fig. 126A and Fig. 126D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 126B and Fig. 126E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 126C and Fig. 126F are photographs of a Grade 2 invasive carcinoma.

[00389] **Figure 127A-127C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 5C6F3 at 1ug/mL. Fig. 127A shows photographs of the tissue micro array. Fig. 127B shows map of the array with abbreviated tissue descriptors. Fig. 127C detailed description of the tissue micro array with non-identifying donor data.

[00390] **Figure 128A-128X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 5C6F3 at 1ug/mL, magnified to 6X and 20X. Fig. 128A and Fig. 128E are adrenal gland. Fig. 128B and Fig. 128F are breast. Fig. 128C and Fig. 128G are fallopian tube. Fig. 128D and Fig. 128H are kidney. Fig. 128I and Fig. 128M are heart muscle. Fig. 128J and Fig. 128N are liver. Fig. 128K and Fig. 128O are lung. Fig. 128L and Fig. 128P are ureter. Fig. 128Q and Fig. 128U are eye. Fig. 128R and Fig. 128V are cerebral cortex. Fig. 128S and Fig. 128W are bone marrow. Fig. 128T and Fig. 128X are skeletal muscle.

[00391] **Figure 129A-129C** shows photographs, array map and description of pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 5C6F3 at 1-20ug/mL. Fig. 129A shows photographs of the tissue micro array. Fig. 129B shows map of the array with



abbreviated tissue descriptors. Fig. 129C detailed description of the tissue micro array with non-identifying donor data.

**[00392]** **Figure 130A-130F** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 5C6F3 at 1ug/mL, magnified to 6X and 20X. Fig. 130A and Fig. 130D are photographs of a Grade 2 adenocarcinoma. Fig. 130B and Fig. 130E are photographs of a Grade 2 adenocarcinoma. Fig. 130C and Fig. 130F are photographs of a Grade 2 adenocarcinoma.

**[00393]** **Figure 131A-131C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 5C6F3 at 1ug/mL. Fig. 131A shows photographs of the tissue micro array. Fig. 131B shows map of the array with abbreviated tissue descriptors. Fig. 131C detailed description of the tissue micro array with non-identifying donor data.

**[00394]** **Figure 132A-132F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 5C6F3 at 1ug/mL, magnified to 6X and 20X. Fig. 132A and Fig. 132D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 132B and Fig. 132E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 132C and Fig. 132F are photographs of a Grade 2 invasive carcinoma.

**[00395]** **Figure 133A-133C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 18B4 at 10ug/mL. Fig. 133A shows photographs of the tissue micro array. Fig. 133B shows map of the array with abbreviated tissue descriptors. Fig. 133C detailed description of the tissue micro array with non-identifying donor data.

**[00396]** **Figure 134A-134X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 18B4 at 10ug/mL, magnified to 6X and 20X. Fig. 134A and Fig. 134E are adrenal gland. Fig. 134B and Fig. 134F are breast. Fig. 134C and Fig. 134G are fallopian tube. Fig. 134D and Fig. 134H are kidney. Fig. 134I and Fig. 134M are heart muscle. Fig. 134J and Fig. 134N are liver. Fig. 134K and Fig. 134O are lung. Fig. 134L and Fig. 134P are ureter. Fig. 134Q and Fig. 134U are eye. Fig. 134R and Fig. 134V are cerebral cortex. Fig. 134S and Fig. 134W are bone marrow. Fig. 134T and Fig. 134X are skeletal muscle.

[00397] **Figure 135A-135C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 18B4 at 10ug/mL. Fig. 135A shows photographs of the tissue micro array. Fig. 135B shows map of the array with abbreviated tissue descriptors. Fig. 135C detailed description of the tissue micro array with non-identifying donor data.

[00398] **Figure 136A-136F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 18B4 at 10ug/mL, magnified to 6X and 20X. Fig. 136A and Fig. 136D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 136B and Fig. 136E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 136C and Fig. 136F are photographs of a Grade 2 invasive ductal carcinoma.

[00399] **Figure 137A-137C** shows photographs, array map and description of esophageal cancer tissue array BC001113 stained with the anti-PSMGFR antibody 18B4 at 10ug/mL. Fig. 137A shows photographs of the tissue micro array. Fig. 137B shows map of the array with abbreviated tissue descriptors. Fig. 137C detailed description of the tissue micro array with non-identifying donor data.

[00400] **Figure 138A-138F** shows photographs of specific tissues from esophageal cancer tissue array BC001113 stained with the anti-PSMGFR antibody 18B4 at 10ug/mL, magnified to 6X and 20X. Fig. 138A and Fig. 138D are photographs of the specimen at position A1. Fig. 138B and Fig. 138E are photographs of the specimen at position A7. Fig. 138C and Fig. 138F are photographs of the specimen at position A8.

[00401] **Figure 139A-139C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 18G12 at 10ug/mL. Fig. 139A shows photographs of the tissue micro array. Fig. 139B shows map of the array with abbreviated tissue descriptors. Fig. 139C detailed description of the tissue micro array with non-identifying donor data.

[00402] **Figure 140A-140X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 18G12 at 10ug/mL, magnified to 6X and 20X. Fig. 140A and Fig. 140E are adrenal gland. Fig. 140B and Fig. 140F are breast. Fig. 140C and Fig. 140G are fallopian tube. Fig. 140D and Fig. 140H are kidney. Fig. 140I and Fig. 140M are heart muscle. Fig. 140J and Fig. 140N are liver. Fig. 140K and Fig. 140O are lung. Fig. 140L and Fig. 140P are ureter. Fig. 140Q and Fig. 140U are eye. Fig. 140R and Fig. 140V

are cerebral cortex. Fig. 140S and Fig. 140W are bone marrow. Fig. 140T and Fig. 140X are skeletal muscle.

**[00403]** **Figure 141A-141C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 18G12 at 15ug/mL. Fig. 141A shows photographs of the tissue micro array. Fig. 141B shows map of the array with abbreviated tissue descriptors. Fig. 141C detailed description of the tissue micro array with non-identifying donor data.

**[00404]** **Figure 142A-142F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 18G12 at 15ug/mL, magnified to 6X and 20X. Fig. 142A and Fig. 142D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 142B and Fig. 142E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 142C and Fig. 142F are photographs of a Grade 2 invasive ductal carcinoma.

**[00405]** **Figure 143A-143C** shows photographs, array map and description of pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 18G12 at 15ug/mL. Fig. 143A shows photographs of the tissue micro array. Fig. 143B shows map of the array with abbreviated tissue descriptors. Fig. 143C detailed description of the tissue micro array with non-identifying donor data.

**[00406]** **Figure 144A-144F** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 18G12 at 15ug/mL, magnified to 6X and 20X. Fig. 144A and Fig. 144D are photographs of a Grade 2 adenocarcinoma. Fig. 144B and Fig. 144E are photographs of a Grade 2 adenocarcinoma. Fig. 144C and Fig. 144F are photographs of a Grade 2-3 adenocarcinoma with lymph node involvement.

**[00407]** **Figure 145A-145C** shows photographs, array map and description of esophageal cancer tissue array BC001113 stained with the anti-PSMGFR antibody 18G12 at 30ug/mL. Fig. 145A shows photographs of the tissue micro array. Fig. 145B shows map of the array with abbreviated tissue descriptors. Fig. 145C detailed description of the tissue micro array with non-identifying donor data.

**[00408]** **Figure 146A-146F** shows photographs of specific tissues from esophageal cancer tissue array BC001113 stained with the anti-PSMGFR antibody 18G12 at 30ug/mL, magnified to 6X and 20X. Fig. 146A and Fig. 146D are photographs of the specimen at position A1. Fig.

146B and Fig. 146E are photographs of the specimen at position A7. Fig. 146C and Fig. 146F are photographs of the specimen at position A8.

**[00409]** **Figure 147A-147C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 25E6 at 5.0ug/mL. Fig. 147A shows photographs of the tissue micro array. Fig. 147B shows map of the array with abbreviated tissue descriptors. Fig. 147C detailed description of the tissue micro array with non-identifying donor data.

**[00410]** **Figure 148A-148X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 25E6 at 5.0ug/mL, magnified to 6X and 20X. Fig. 148A and Fig. 148E are adrenal gland. Fig. 148B and Fig. 148F are breast. Fig. 148C and Fig. 148G are fallopian tube. Fig. 148D and Fig. 148H are kidney. Fig. 148I and Fig. 148M are heart muscle. Fig. 148J and Fig. 148N are liver. Fig. 148K and Fig. 148O are lung. Fig. 148L and Fig. 148P are ureter. Fig. 148Q and Fig. 148U are eye. Fig. 148R and Fig. 148V are cerebral cortex. Fig. 148S and Fig. 148W are bone marrow. Fig. 148T and Fig. 148X are skeletal muscle.

**[00411]** **Figure 149A-149C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 25E6 at 5.0ug/mL. Fig. 149A shows photographs of the tissue micro array. Fig. 149B shows map of the array with abbreviated tissue descriptors. Fig. 149C detailed description of the tissue micro array with non-identifying donor data.

**[00412]** **Figure 150A-150F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 25E6 at 5.0ug/mL, magnified to 6X and 20X. Fig. 150A and Fig. 150D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 150B and Fig. 150E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 150C and Fig. 150F are photographs of a Grade 2 invasive ductal carcinoma.

**[00413]** **Figure 151A-151C** shows photographs, array map and description of pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 25E6 at 5.0ug/mL. Fig. 151A shows photographs of the tissue micro array. Fig. 151B shows map of the array with abbreviated tissue descriptors. Fig. 151C detailed description of the tissue micro array with non-identifying donor data.

**[00414]** **Figure 152A-152F** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 25E6 at 5.0ug/mL, magnified to 6X and 20X. Fig. 152A and Fig. 152D are photographs of a Grade 2 adenocarcinoma. Fig. 152B and Fig. 152E are photographs of a Grade 1 adenocarcinoma. Fig. 152C and Fig. 152F are photographs of a Grade 1 adenocarcinoma.

**[00415]** **Figure 153A-153C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 28F9 at 15.0ug/mL. Fig. 153A shows photographs of the tissue micro array. Fig. 153B shows map of the array with abbreviated tissue descriptors. Fig. 153C detailed description of the tissue micro array with non-identifying donor data.

**[00416]** **Figure 154A-154X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 28F9 at 15.0ug/mL, magnified to 6X and 20X. Fig. 154A and Fig. 154E are adrenal gland. Fig. 154B and Fig. 154F are breast. Fig. 154C and Fig. 154G are fallopian tube. Fig. 154D and Fig. 154H are kidney. Fig. 154I and Fig. 154M are heart muscle. Fig. 154J and Fig. 154N are liver. Fig. 154K and Fig. 154O are lung. Fig. 154L and Fig. 154P are ureter. Fig. 154Q and Fig. 154U are eye. Fig. 154R and Fig. 154V are cerebral cortex. Fig. 154S and Fig. 154W are bone marrow. Fig. 154T and Fig. 154X are skeletal muscle.

**[00417]** **Figure 155A-155C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 28F9 at 15.0ug/mL. Fig. 155A shows photographs of the tissue micro array. Fig. 155B shows map of the array with abbreviated tissue descriptors. Fig. 155C detailed description of the tissue micro array with non-identifying donor data.

**[00418]** **Figure 156A-156F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 28F9 at 15.0ug/mL, magnified to 6X and 20X. Fig. 156A and Fig. 156D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 156B and Fig. 156E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 156C and Fig. 156F are photographs of a Grade 2 invasive ductal carcinoma.

**[00419]** **Figure 157A-157C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the N+20/C-27 antibody 1E4 at 7.5ug/mL. Fig. 157A shows photographs of the tissue micro array. Fig. 157B shows map of the array with abbreviated tissue

descriptors. Fig. 157C detailed description of the tissue micro array with non-identifying donor data.

**[00420]** **Figure 158A-158X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the N+20/C-27 antibody 1E4 at 7.5ug/mL, magnified to 6X and 20X. Fig. 158A and Fig. 158E are adrenal gland. Fig. 158B and Fig. 158F are breast. Fig. 158C and Fig. 158G are fallopian tube. Fig. 158D and Fig. 158H are kidney. Fig. 158I and Fig. 158M are heart muscle. Fig. 158J and Fig. 158N are liver. Fig. 158K and Fig. 158O are lung. Fig. 158L and Fig. 158P are ureter. Fig. 158Q and Fig. 158U are eye. Fig. 158R and Fig. 158V are cerebral cortex. Fig. 158S and Fig. 158W are bone marrow. Fig. 158T and Fig. 158X are skeletal muscle.

**[00421]** **Figure 159A-159C** shows photographs, array map and description of breast cancer tissue array BR1007 stained with the N+20/C-27 antibody 1E4 at 10.0ug/mL. Fig. 159A shows photographs of the tissue micro array. Fig. 159B shows map of the array with abbreviated tissue descriptors. Fig. 159C detailed description of the tissue micro array with non-identifying donor data.

**[00422]** **Figure 160A-160F** shows photographs of specific tissues from breast cancer tissue array BR1007 stained with the N+20/C-27 antibody 1E4 at 10.0ug/mL, magnified to 6X and 20X. Fig. 160A and Fig. 160D are photographs of a Grade 2 invasive ductal carcinoma with positive lymph nodes. Fig. 160B and Fig. 160E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 160C and Fig. 160F are photographs of a Grade 2 invasive ductal carcinoma.

**[00423]** **Figure 161A-161C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the N+20/C-27 antibody 29H1 at 0.5ug/mL. Fig. 161A shows photographs of the tissue micro array. Fig. 161B shows map of the array with abbreviated tissue descriptors. Fig. 161C detailed description of the tissue micro array with non-identifying donor data.

**[00424]** **Figure 162A-162X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the N+20/C-27 antibody 29H1 at 0.5ug/mL, magnified to 6X and 20X. Fig. 162A and Fig. 162E are adrenal gland. Fig. 162B and Fig. 162F are breast. Fig. 162C and Fig. 162G are fallopian tube. Fig. 162D and Fig. 162H are kidney. Fig. 162I and Fig. 162M are heart muscle. Fig. 162J and Fig. 162N are liver. Fig. 162K and Fig. 162O are lung. Fig. 162L and Fig. 162P are ureter. Fig. 162Q and Fig. 162U are eye. Fig. 162R and Fig. 162V are

cerebral cortex. Fig. 162S and Fig. 162W are bone marrow. Fig. 162T and Fig. 162X are skeletal muscle.

**[00425]** **Figure 163A-163C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the N+20/C-27 antibody 29H1 at 0.5ug/mL. Fig. 163A shows photographs of the tissue micro array. Fig. 163B shows map of the array with abbreviated tissue descriptors. Fig. 163C detailed description of the tissue micro array with non-identifying donor data.

**[00426]** **Figure 164A-164F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the N+20/C-27 antibody 29H1 at 0.5ug/mL, magnified to 6X and 20X. Fig. 164A and Fig. 164D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 164B and Fig. 164E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 164C and Fig. 164F are photographs of a Grade 2 invasive ductal carcinoma.

**[00427]** **Figure 165A-165C** shows photographs, array map and description of pancreatic cancer tissue array PA1003 stained with the N+20/C-27 antibody 29H1 at 0.5ug/mL. Fig. 165A shows photographs of the tissue micro array. Fig. 165B shows map of the array with abbreviated tissue descriptors. Fig. 165C detailed description of the tissue micro array with non-identifying donor data.

**[00428]** **Figure 166A-166F** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the N+20/C-27 antibody 29H1 at 0.5ug/mL, magnified to 6X and 20X. Fig. 166A and Fig. 166D are photographs of a Grade 2 adenocarcinoma. Fig. 166B and Fig. 166E are photographs of a Grade 2 adenocarcinoma. Fig. 166C and Fig. 166F are photographs of a Grade 3 adenocarcinoma.

**[00429]** **Figure 167A-167C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the N+20/C-27 antibody 31A1 at 0.5ug/mL. Fig. 167A shows photographs of the tissue micro array. Fig. 167B shows map of the array with abbreviated tissue descriptors. Fig. 167C detailed description of the tissue micro array with non-identifying donor data.

**[00430]** **Figure 168A-168X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the N+20/C-27 antibody 31A1 at 0.5ug/mL, magnified to 6X and 20X. Fig. 168A and Fig. 168E are adrenal gland. Fig. 168B and Fig. 168F are breast. Fig. 168C and Fig. 168G are fallopian tube. Fig. 168D and Fig. 168H are kidney. Fig. 168I and Fig. 168M

are heart muscle. Fig. 168J and Fig. 168N are liver. Fig. 168K and Fig. 168O are lung. Fig. 168L and Fig. 168P are ureter. Fig. 168Q and Fig. 168U are eye. Fig. 168R and Fig. 168V are cerebral cortex. Fig. 168S and Fig. 168W are bone marrow. Fig. 168T and Fig. 168X are skeletal muscle.

**[00431]** **Figure 169A-169C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the N+20/C-27 antibody 31A1 at 0.5ug/mL. Fig. 169A shows photographs of the tissue micro array. Fig. 169B shows map of the array with abbreviated tissue descriptors. Fig. 169C detailed description of the tissue micro array with non-identifying donor data.

**[00432]** **Figure 170A-170F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the N+20/C-27 antibody 31A1 at 0.5ug/mL, magnified to 6X and 20X. Fig. 170A and Fig. 170D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 170B and Fig. 170E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 170C and Fig. 170F are photographs of a Grade 2 invasive ductal carcinoma.

**[00433]** **Figure 171A-171C** shows photographs, array map and description of pancreatic cancer tissue array PA1003 stained with the N+20/C-27 antibody 31A1 at 0.5ug/mL. Fig. 171A shows photographs of the tissue micro array. Fig. 171B shows map of the array with abbreviated tissue descriptors. Fig. 171C detailed description of the tissue micro array with non-identifying donor data.

**[00434]** **Figure 172A-172F** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the N+20/C-27 antibody 31A1 at 0.5ug/mL, magnified to 6X and 20X. Fig. 172A and Fig. 172D are photographs of a Grade 1 adenocarcinoma. Fig. 172B and Fig. 172E are photographs of a Grade 2 adenocarcinoma. Fig. 172C and Fig. 172F are photographs of a Grade 3 adenocarcinoma.

**[00435]** **Figure 173A-173C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the N+20/C-27 antibody 32C1 at 0.25ug/mL. Fig. 173A shows photographs of the tissue micro array. Fig. 173B shows map of the array with abbreviated tissue descriptors. Fig. 173C detailed description of the tissue micro array with non-identifying donor data.

**[00436]** **Figure 174A-174X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the N+20/C-27 antibody 32C1 at 0.25ug/mL, magnified to 6X and 20X.



Fig. 174A and Fig. 174E are adrenal gland. Fig. 174B and Fig. 174F are breast. Fig. 174C and Fig. 174G are fallopian tube. Fig. 174D and Fig. 174H are kidney. Fig. 174I and Fig. 174M are heart muscle. Fig. 174J and Fig. 174N are liver. Fig. 174K and Fig. 174O are lung. Fig. 174L and Fig. 174P are ureter. Fig. 174Q and Fig. 174U are eye. Fig. 174R and Fig. 174V are cerebral cortex. Fig. 174S and Fig. 174W are bone marrow. Fig. 174T and Fig. 174X are skeletal muscle.

**[00437]** **Figure 175A-175C** shows photographs, array map and description of breast cancer tissue array 1141 stained with the N+20/C-27 antibody 32C1 at 5.0ug/mL. Fig. 175A shows photographs of the tissue micro array. Fig. 175B shows map of the array with abbreviated tissue descriptors. Fig. 175C detailed description of the tissue micro array with non-identifying donor data.

**[00438]** **Figure 176A-176F** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the N+20/C-27 antibody 32C1 at 5.0ug/mL, magnified to 6X and 20X. Fig. 176A and Fig. 176D are photographs of a Grade 2 invasive ductal carcinoma. Fig. 176B and Fig. 176E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 176C and Fig. 176F are photographs of a Grade 2 invasive ductal carcinoma.

**[00439]** **Figure 177A-177C** shows photographs, array map and description of esophageal cancer tissue array ES1001 stained with the N+20/C-27 antibody 32C1 at 1.0ug/mL. Fig. 177A shows photographs of the tissue micro array. Fig. 177B shows map of the array with abbreviated tissue descriptors. Fig. 177C detailed description of the tissue micro array with non-identifying donor data.

**[00440]** **Figure 178A-178F** shows photographs of specific tissues from esophageal cancer tissue array BC001113 stained with the N+20/C-27 antibody 32C1 at 1.0ug/mL, magnified to 6X and 20X. Fig. 178A and Fig. 178D are photographs of a squamous cell carcinoma. Fig. 178B and Fig. 178E are photographs of an adenocarcinoma. Fig. 178C and Fig. 178F are photographs of a squamous cell carcinoma.

**[00441]** **Figure 179A-179C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the N+20/C-27 antibody 45C11 at 12.5ug/mL. Fig. 179A shows photographs of the tissue micro array. Fig. 179B shows map of the array with abbreviated tissue descriptors. Fig. 179C detailed description of the tissue micro array with non-identifying donor data.

**[00442]** **Figure 180A-180X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the N+20/C-27 antibody 45C11 at 12.5ug/mL, magnified to 6X and 20X. Fig. 180A and Fig. 180E are adrenal gland. Fig. 180B and Fig. 180F are breast. Fig. 180C and Fig. 180G are fallopian tube. Fig. 180D and Fig. 180H are kidney. Fig. 180I and Fig. 180M are heart muscle. Fig. 180J and Fig. 180N are liver. Fig. 180K and Fig. 180O are lung. Fig. 180L and Fig. 180P are ureter. Fig. 180Q and Fig. 180U are eye. Fig. 180R and Fig. 180V are cerebral cortex. Fig. 180S and Fig. 180W are bone marrow. Fig. 180T and Fig. 180X are skeletal muscle.

**[00443]** **Figure 181A-181C** shows photographs, array map and description of breast cancer tissue array BR1007 stained with the N+20/C-27 antibody 45C11 at 10.0ug/mL. Fig. 181A shows photographs of the tissue micro array. Fig. 181B shows map of the array with abbreviated tissue descriptors. Fig. 181C detailed description of the tissue micro array with non-identifying donor data.

**[00444]** **Figure 182A-182F** shows photographs of specific tissues from breast cancer tissue array BR1007 stained with the N+20/C-27 antibody 45C11 at 10.0ug/mL, magnified to 6X and 20X. Fig. 182A and Fig. 182D are photographs of a Grade 2 invasive ductal carcinoma with positive lymph nodes. Fig. 182B and Fig. 182E are photographs of a Grade 2 invasive ductal carcinoma. Fig. 182C and Fig. 182F are photographs of a Grade 2 invasive ductal carcinoma.

**[00445]** **Figure 183A-183C** shows photographs, array map and description of pancreatic cancer tissue array PA805c stained with the N+20/C-27 antibody 45C11 at 12.5ug/mL. Fig. 183A shows photographs of the tissue micro array. Fig. 183B shows map of the array with abbreviated tissue descriptors. Fig. 183C detailed description of the tissue micro array with non-identifying donor data.

**[00446]** **Figure 184A-184F** shows photographs of specific tissues from pancreatic cancer tissue array PA805c stained with the N+20/C-27 antibody 45C11 at 12.5ug/mL, magnified to 6X and 20X. Fig. 184A and Fig. 184D are photographs of a Grade 2 papillary adenocarcinoma. Fig. 184B and Fig. 184E are photographs of a Grade 2-3 ductal carcinoma. Fig. 184C and Fig. 184F are photographs of a Grade 3 invasive adenocarcinoma.

**[00447]** **Figure 185A-185C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the N+9/C-9 antibody 3C5 at 10.0ug/mL. Fig. 185A shows photographs of the tissue micro array. Fig. 185B shows map of the array with abbreviated tissue

descriptors. Fig. 185C detailed description of the tissue micro array with non-identifying donor data.

**[00448] Figure 186A-186X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the N+9/C-9 antibody 3C5 at 10.0ug/mL, magnified to 6X and 20X. Fig. 186A and Fig. 186E are adrenal gland. Fig. 186B and Fig. 186F are breast. Fig. 186C and Fig. 186G are fallopian tube. Fig. 186D and Fig. 186H are kidney. Fig. 186I and Fig. 186M are heart muscle. Fig. 186J and Fig. 186N are liver. Fig. 186K and Fig. 186O are lung. Fig. 186L and Fig. 186P are ureter. Fig. 186Q and Fig. 186U are eye. Fig. 186R and Fig. 186V are cerebral cortex. Fig. 186S and Fig. 186W are bone marrow. Fig. 186T and Fig. 186X are skeletal muscle.

**[00449] Figure 187A-187C** shows photographs, array map and description of pancreatic cancer tissue array PA1003 stained with the N+9/C-9 antibody 3C5 at 10.0ug/mL. Fig. 187A shows photographs of the tissue micro array. Fig. 187B shows map of the array with abbreviated tissue descriptors. Fig. 187C detailed description of the tissue micro array with non-identifying donor data.

**[00450] Figure 188A-188F** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the N+9/C-9 antibody 3C5 at 10.0ug/mL, magnified to 6X and 20X. Fig. 188A and Fig. 188D are photographs of a Grade 2 adenocarcinoma. Fig. 188B and Fig. 188E are photographs of a Grade 2 adenocarcinoma. Fig. 188C and Fig. 188F are photographs of a Grade 2-3 adenocarcinoma with lymph node involvement.

**[00451] Figure 189A-189C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the N+9/C-9 antibody 8A9 at 15.0ug/mL. Fig. 189A shows photographs of the tissue micro array. Fig. 189B shows map of the array with abbreviated tissue descriptors. Fig. 189C detailed description of the tissue micro array with non-identifying donor data.

**[00452] Figure 190A-190X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the N+9/C-9 antibody 8A9 at 15.0ug/mL, magnified to 6X and 20X. Fig. 190A and Fig. 190E are adrenal gland. Fig. 190B and Fig. 190F are breast. Fig. 190C and Fig. 190G are fallopian tube. Fig. 190D and Fig. 190H are kidney. Fig. 190I and Fig. 190M are heart muscle. Fig. 190J and Fig. 190N are liver. Fig. 190K and Fig. 190O are lung. Fig. 190L and Fig. 190P are ureter. Fig. 190Q and Fig. 190U are eye. Fig. 190R and Fig. 190V are cerebral cortex. Fig. 190S and Fig. 190W are bone marrow. Fig. 190T and Fig. 190X are skeletal muscle.

**[00453]** **Figure 191A-191C** shows photographs, array map and description of pancreatic cancer tissue array PA1003 stained with the N+9/C-9 antibody 8A9 at 15.0ug/mL. Fig. 191A shows photographs of the tissue micro array. Fig. 191B shows map of the array with abbreviated tissue descriptors. Fig. 191C detailed description of the tissue micro array with non-identifying donor data.

**[00454]** **Figure 192A-192F** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the N+9/C-9 antibody 8A9 at 15.0ug/mL, magnified to 6X and 20X. Fig. 192A and Fig. 192D are photographs of a Grade 2 adenocarcinoma. Fig. 192B and Fig. 192E are photographs of a Grade 2 adenocarcinoma. Fig. 192C and Fig. 192F are photographs of a Grade 2 adenocarcinoma.

**[00455]** **Figure 193A-193C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the N+9/C-9 antibody 17H6 at 30.0ug/mL. Fig. 193A shows photographs of the tissue micro array. Fig. 193B shows map of the array with abbreviated tissue descriptors. Fig. 193C detailed description of the tissue micro array with non-identifying donor data.

**[00456]** **Figure 194A-194X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the N+9/C-9 antibody 17H6 at 30.0ug/mL, magnified to 6X and 20X. Fig. 194A and Fig. 194E are adrenal gland. Fig. 194B and Fig. 194F are breast. Fig. 194C and Fig. 194G are fallopian tube. Fig. 194D and Fig. 194H are kidney. Fig. 194I and Fig. 194M are heart muscle. Fig. 194J and Fig. 194N are liver. Fig. 194K and Fig. 194O are lung. Fig. 194L and Fig. 194P are ureter. Fig. 194Q and Fig. 194U are eye. Fig. 194R and Fig. 194V are cerebral cortex. Fig. 194S and Fig. 194W are bone marrow. Fig. 194T and Fig. 194X are skeletal muscle.

**[00457]** **Figure 195A-195C** shows photographs, array map and description of pancreatic cancer tissue array PA805c stained with the N+9/C-9 antibody 17H6 at 30.0ug/mL. Fig. 195A shows photographs of the tissue micro array. Fig. 195B shows map of the array with abbreviated tissue descriptors. Fig. 195C detailed description of the tissue micro array with non-identifying donor data.

**[00458]** **Figure 196A-196F** shows photographs of specific tissues from pancreatic cancer tissue array PA805c stained with the N+9/C-9 antibody 17H6 at 30.0ug/mL, magnified to 6X and 20X. Fig. 196A and Fig. 196D are photographs of a Grade 2 papillary adenocarcinoma. Fig.

196B and Fig. 196E are photographs of a Grade 2-3 ductal carcinoma with lymph node involvement. Fig. 196C and Fig. 196F are photographs of a Grade 3 invasive adenocarcinoma.

**[00459]** **Figure 197A-197C** shows photographs, array map and description of FDA normal tissue array 1021 stained with the N+9/C-9 antibody 39H5 at 5.0ug/mL. Fig. 197A shows photographs of the tissue micro array. Fig. 197B shows map of the array with abbreviated tissue descriptors. Fig. 197C detailed description of the tissue micro array with non-identifying donor data.

**[00460]** **Figure 198A-198X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the N+9/C-9 antibody 39H5 at 5.0ug/mL, magnified to 6X and 20X. Fig. 198A and Fig. 198E are adrenal gland. Fig. 198B and Fig. 198F are breast. Fig. 198C and Fig. 198G are fallopian tube. Fig. 198D and Fig. 198H are kidney. Fig. 198I and Fig. 198M are heart muscle. Fig. 198J and Fig. 198N are liver. Fig. 198K and Fig. 198O are lung. Fig. 198L and Fig. 198P are ureter. Fig. 198Q and Fig. 198U are eye. Fig. 198R and Fig. 198V are cerebral cortex. Fig. 198S and Fig. 198W are bone marrow. Fig. 198T and Fig. 198X are skeletal muscle.

**[00461]** **Figure 199A-199C** shows photographs, array map and description of pancreatic cancer tissue array PA1003 stained with the N+9/C-9 antibody 39H5 at 5.0ug/mL. Fig. 199A shows photographs of the tissue micro array. Fig. 199B shows map of the array with abbreviated tissue descriptors. Fig. 199C detailed description of the tissue micro array with non-identifying donor data.

**[00462]** **Figure 200A-200F** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the N+9/C-9 antibody 39H5 at 5.0ug/mL, magnified to 6X and 20X. Fig. 200A and Fig. 200D are photographs of a Grade 2 adenocarcinoma. Fig. 200B and Fig. 200E are photographs of a Grade 2 adenocarcinoma. Fig. 200C and Fig. 200F are photographs of a Grade 2 adenocarcinoma.

**[00463]** **Figure 201A-201C** show graphs of ELISA assays to determine the binding of another set of antibodies generated by immunizing animals with the PSMGFR peptide. Fig. 201A shows binding to the PSMGFR peptide. Fig. 201B shows binding to the N-10 peptide. Fig. 201C shows binding to the C-10 peptide. As can be seen, none of the antibodies bound to the C-10 peptide. F3, B12, B2, B7, B9, 8C7F3 and H11 all bound to the PSMGFR peptide and to the N-10 peptide.

**[00464]** **Figure 202A-202C** shows photographs of pancreatic cancer tissue array PA1003 that has been stained with monoclonal antibody 1E4, monoclonal antibody 18B4 or the polyclonal

anti-PSMGFR antibody SDIX. 18B4 binds to the GTINVHDTVET epitope at the most N-terminal portion of the PSMGFR peptide, while the 1E4 antibody binds to the QFNQYKTEA epitope that is immediately adjacent and C-terminal to the 18B4 epitope.

**[00465]** **Figure 203A-203F** shows magnified images of the tissue specimen at position A2 of the pancreatic cancer array PA1003. Fig. 203A and Fig. 203B show the specimen stained with antibody 1E4. Fig. 203C and Fig. 203D show the specimen stained with antibody 18B4. Fig. 203E and Fig. 203F show the specimen stained with polyclonal antibody SDIX.

**[00466]** **Figure 204A-204D** shows magnified images of the tissue specimen at position D4 of the pancreatic array PA1003. Fig. 204A and Fig. 204B show the specimen stained with antibody 18B4. Fig. 204C and Fig. 204D show the specimen stained with polyclonal antibody SDIX.

**[00467]** **Figure 205A-205D** shows magnified images of the tissue specimen at position E1 of the pancreatic cancer array PA1003. Fig. 205A and Fig. 205B show the specimen stained with antibody 18B4. Fig. 205C and Fig. 205D show the specimen stained with polyclonal antibody SDIX.

**[00468]** **Figure 206A-206D** shows magnified images of the tissue specimen at position C3 of the pancreatic cancer array PA1003. Fig. 206A and Fig. 206B show the specimen stained with antibody 1E4. Fig. 206C and Fig. 206D show the specimen stained with polyclonal antibody SDIX.

**[00469]** **Figure 207A-207D** shows magnified images of the tissue specimen at position D1 of the pancreatic cancer array PA1003. Fig. 207A and Fig. 207B show the specimen stained with antibody 1E4. Fig. 207C and Fig. 207D show the specimen stained with polyclonal antibody SDIX.

**[00470]** **Figure 208A-208C** shows photographs of the pancreatic cancer array PA1003. Fig. 208A shows the specimen stained with polyclonal antibody SDIX. Fig. 208B shows the specimen stained with antibody 20A10. Fig. 208C shows the specimen stained with antibody 29H1.

**[00471]** **Figure 209A-209D** shows photographs of the esophageal cancer array ES1001 stained with various antibodies. Fig. 209A shows the array stained with polyclonal antibody SDIX. Fig. 209B shows the array stained with antibody 20A10. Fig. 209C shows the array stained with antibody 29H1. Fig. 209D shows the array stained with antibody 31A1.

[00472] **Figure 210A-210C** shows photographs of the pancreatic cancer array PA1003 stained with various antibodies. Fig. 210A shows the array stained with polyclonal antibody SDIX. Fig. 210B shows the array stained with antibody 20A10. Fig. 210C shows the array stained with antibody 29H1.

[00473] **Figure 211A- 211C** show graphs of an ELISA experiment measuring the amount of IL-18 secreted into the condition media of MUC1\* positive cancer cells co-cultured with huMNC2-CAR44 T cells wherein the cells also bear an NFAT inducible IL-18. Fig. 211A shows the graph of IL-18 secreted into the supernatant of T47D breast cancer cells co-cultured with untransduced human T cells. Fig. 211B shows the graph of IL-18 secreted into the supernatant of T47D breast cancer cells co-cultured with huMNC2-CAR44 T cells that also bore an NFAT inducible IL-18 gene inserted into a portion of the Foxp3 enhancer. Fig. 211C shows the graph of IL-18 secreted into the supernatant of T47D breast cancer cells co-cultured with huMNC2-CAR44 T cells that also bore an NFAT inducible IL-18 gene inserted into a portion of the IL-2 enhancer.

[00474] **Figure 212A- 212X** shows photographs of T47D breast cancer cells (red) doped with varying percentages of T47D cells engineered to express more MUC1\* (green). The target cancer cells have been co-cultured with huMNC2-CAR44 T cells with NFAT inducible IL-18 wherein the IL-18 gene has been inserted into either the Foxp3 enhancer/promoter or the IL-2 enhancer/promoter. Fig. 212A-212C, 212I-212K, and 212Q-212S show the cancer cells co-cultured with untransduced T cells. Fig. 212D-212F, 212L-212N, and 212T-212V show the cancer cells co-cultured with huMNC2-CAR44 T cells with the NFAT inducible IL-18 gene inserted into the Foxp3 enhancer/promoter. Fig. 212G-212H, 212O-212P, and 212W-212X show the cancer cells co-cultured with huMNC2-CAR44 T cells with the NFAT inducible IL-18 gene inserted into the IL-2 enhancer/promoter.

[00475] **Figure 213A- 213B** shows graphs of ELISA experiments in which levels of IL-18 secreted into the conditioned media are measured for huMNC1-CAR44 T cells with NFAT inducible IL-18 gene, inserted into the Foxp3 enhancer or promoter, co-cultured with either MUC1\* positive cancer cells or MUC1 negative non-cancerous cells. Fig. 213A shows IL-18 secretion from huMNC2-CAR44 T cells with NFAT inducible IL-18 in co-culture with T47D breast cancer cells where the population has been doped with 5%, 10% or 30% T47D cells that had been transfected with even more MUC1\*. Fig. **213B** shows IL-18 secretion from huMNC2-

CAR44 T cells with NFAT inducible IL-18 in co-culture with non-cancerous, MUC1 negative HEK293 cells where the cell population has been doped with 5%, 10% or 30% T47D cells that had been transfected with more MUC1\*.

**[00476]** **Figure 214A-214X** shows photographs of T47D breast cancer cells (red) or non-cancerous HEK293 cells (also red), where both cell types have been doped with varying percentages of T47D cells engineered to express more MUC1\* (green). These target cancer cells have been co-cultured with huMNC2-CAR44 T cells with NFAT inducible IL-18 wherein the IL-18 gene has been inserted into the Foxp3 enhancer/promoter. Fig. 214A-214F shows either T47D cells or HEK293 cells that have not been doped with T47D cells engineered to express high MUC1\* density. Fig. 214G-214L shows either T47D cells or HEK293 cells that have been doped with 5% T47D cells engineered to express high MUC1\* density. Fig. 214M-214R shows either T47D cells or HEK293 cells that have been doped with 10% T47D cells engineered to express high MUC1\* density. Fig. 214S-214X shows either T47D cells or HEK293 cells that have been doped with 30% T47D cells engineered to express high MUC1\* density. Fig. 214A-B, G-H, M-N, and S-T show T47D breast cancer cells. Fig. 214C-F, I-L, O-R, and U-X show HEK293 cells. As can be seen in the figures, the induced secretion of IL-18 resulted in low MUC1\* density T47D cells being killed but did not induce non-specific killing of the MUC1\* negative HEK293 cells.

**[00477]** **Figure 215A-215C** shows the consensus sequences of the heavy chain CDRs wherein the consensus sequences were generated for each group of antibodies that bound to the same epitope in the PSMGFR and N-terminally extended PSMGFR peptide. Fig. 215A shows consensus sequences for heavy chain CDR1. Fig. 215B shows consensus sequences for heavy chain CDR2. Fig. 215C shows consensus sequences for heavy chain CDR3.

**[00478]** **Figure 216A-216C** shows the consensus sequences of the light chain CDRs wherein the consensus sequences were generated for each group of antibodies that bound to the same epitope in the PSMGFR and N-terminally extended PSMGFR peptide. Fig. 216A shows consensus sequences for light chain CDR1. Fig. 216B shows consensus sequences for light chain CDR2. Fig. 216C shows consensus sequences for light chain CDR3.



## DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENTS

[00479] In the present application, “a” and “an” are used to refer to both single and a plurality of objects.

[00480] As used herein, occasionally, in short hand, a polypeptide is indicated as being “transduced or transfected” into a cell. In these occurrences, it is understood that the nucleic acid encoding the polypeptide sequence is transduced or transfected into the cell, as it is an impossibility that a polypeptide could be transduced or transfected into a cell.

[00481] As used herein, occasionally when referring to number of cells injected into an animal or otherwise contextually wherein the number of cells is referred to, “M” refers to millions, and “K” refers to thousands.

[00482] As used herein, interchangeable designations for various monoclonal antibodies are used, such as, “MN-C2”, which is interchangeable with “C2”, “Min-C2” and “MNC2”; “MN-E6”, which is interchangeable with “E6”, “Min-E6” and “MNE6”; “MN-C3”, which is interchangeable with “C3”, “Min-C3” and “MNC3”; and “MN-C8”, which is interchangeable with “C8”, “Min-C8” and “MNC8”. The monoclonal antibodies provided herein follow the same convention.

[00483] As used herein, “h” or “hu” placed before an antibody construct is short-hand for humanized.

[00484] As used herein, the term “antibody-like” means a molecule that may be engineered such that it contains portions of antibodies but is not an antibody that would naturally occur in nature. Examples include but are not limited to CAR (chimeric antigen receptor) T cell technology and the Ylanthia<sup>®</sup> technology. The CAR technology uses an antibody epitope fused to a portion of a T cell so that the body’s immune system is directed to attack a specific target protein or cell. The Ylanthia<sup>®</sup> technology consists of an “antibody-like” library that is a collection of synthetic human Fabs that are then screened for binding to peptide epitopes from target proteins. The selected Fab regions can then be engineered into a scaffold or framework so that they resemble antibodies.

[00485] As used herein, “PSMGFR” is abbreviation for Primary Sequence of the MUC1 Growth Factor Receptor which is identified by SEQ ID NO:2, and thus is not to be confused with a six amino acid sequence. “PSMGFR peptide” or “PSMGFR region” refers to a peptide or

region that incorporates the Primary Sequence of the MUC1 Growth Factor Receptor (SEQ ID NO:2).

**[00486]** As used herein, the “MUC1\*” extra cellular domain is defined primarily by the PSMGFR sequence (GTINVHDTVETQFNQYKTEAASRYNLTISDVSVSDVPFPFSAQSGA (SEQ ID NO:2)). Because the exact site of MUC1 cleavage depends on the enzyme that clips it, and that the cleavage enzyme varies depending on cell type, tissue type or the time in the evolution of the cell, the exact sequence of the MUC1\* extra cellular domain may vary at the N-terminus.

**[00487]** Other clipped amino acid sequences may include SNIKFRPGSVVVQLTLAFREGTINVHDTVETQFNQYKTEAASRY (SEQ ID NO:620); or SVVVQLTLAFREGTINVHDTVETQFNQYKTEAASRY (SEQ ID NO:621).

**[00488]** As used herein, the term “PSMGFR” is an acronym for Primary Sequence of MUC1 Growth Factor Receptor as set forth as GTINVHDTVETQFNQYKTEAASRYNLTISDVSVSDVPFPFSAQSGA (SEQ ID NO:2). In this regard, the “N-number” as in “N-10 PSMGFR” or simply “N-10”, “N-15 PSMGFR” or simply “N-15”, or “N-20 PSMGFR” or simply “N-20” refers to the number of amino acid residues that have been deleted at the N-terminal end of PSMGFR. Likewise “C-number” as in “C-10 PSMGFR” or simply “C-10”, “C-15 PSMGFR” or simply “C-15”, or “C-20 PSMGFR” or simply “C-20” refers to the number of amino acid residues that have been deleted at the C-terminal end of PSMGFR. A mixture of deletions and additions is also possible. For instance, N+20/C-27 refers to a peptide fragment of wild-type MUC1 in which 20 amino acids are added to the PSMGFR at the N-terminus and 27 amino acids are deleted from the C-terminus.

**[00489]** As used herein, the “extracellular domain of MUC1\*” refers to the extracellular portion of a MUC1 protein that is devoid of the tandem repeat domain. In most cases, MUC1\* is a cleavage product wherein the MUC1\* portion consists of a short extracellular domain devoid of tandem repeats, a transmembrane domain and a cytoplasmic tail. The precise location of cleavage of MUC1 is not known perhaps because it appears that it can be cleaved by more than one enzyme. The extracellular domain of MUC1\* will include most of the PSMGFR sequence but may have an additional 10-20 N-terminal amino acids.

**[00490]** As used herein “sequence identity” means homology in sequence of a particular polypeptide or nucleic acid to a reference sequence of nucleic acid or amino acid such that the

function of the homologous peptide is the same as the reference peptide or nucleic acid. Such homology can be so close with the reference peptide such that at times the two sequences may be 90%, 95% or 98% identical yet possess the same function in binding or other biological activities.

**[00491]** As used herein, “MUC1 positive” cell refers to a cell that expresses a gene for MUC1, MUC1-Y or MUC1-Z or other MUC1 variant.

**[00492]** As used herein, “MUC1 negative” cell refers to a cell that does not express a gene for MUC1.

**[00493]** As used herein, “MUC1\* positive” cell refers to a cell that expresses a gene for MUC1, wherein that gene’s expressed protein is a transmembrane protein that is devoid of tandem repeats, which may be a consequence of post-translational modification, cleavage, alternative splicing, or transfecting or transducing a cell with a MUC1 protein that is devoid of tandem repeats.

**[00494]** As used herein, “MUC1\* negative” cell refers to a cell that may or may not express a gene for MUC1 but does not express a MUC1 transmembrane protein that is devoid of tandem repeats.

**[00495]** As used herein, “MUC1 positive” cancer cell refers to a cancer cell that overexpresses the gene for MUC1, expresses MUC1 in an aberrant pattern, wherein its expression is not restricted to the apical border and/or expresses a MUC1 that is devoid of tandem repeats.

**[00496]** As used herein, “MUC1 negative” cancer cell refers to a cancer cell that may or may not express a gene for MUC1 but does not overexpress MUC1 or does not overexpress a MUC1 transmembrane protein that is devoid of tandem repeats.

**[00497]** As used herein, “MUC1\* positive” cancer cell refers to a cancer cell that overexpresses a MUC1 transmembrane protein that is devoid of tandem repeats.

**[00498]** As used herein, “MUC1\* negative” cancer cell refers to a cancer cell that may or may not express a gene for MUC1 but does not overexpress a MUC1 transmembrane protein that is devoid of tandem repeats.

**[00499]** As used herein “conformational epitope” refers to a peptide sequence that is required to be present in a specific three-dimensional structure or conformation for an antibody to bind. However the antibody binds when the peptide sequence is in the three-dimensional structure or conformation and is not bound when linear. A common technique for determining whether an

antibody binds to a linear stretch or a conformational epitope is to use the antibody to probe a denaturing Western blot. Traveling through a denaturing gel linearizes proteins and peptides. Antibodies that do not work in a denaturing Western but do recognize the native target, for example expressed on an intact cell, are determined to recognize a conformational epitope. As used herein, the antibody may or may not actually bind to the “conformational epitope”, however the presence of the “conformational epitope” sequence is required to render a three dimensional structure so that the MUC1\* region on cancer cells is able to be bound by the antibody that is specific for cancer treatment. Thus, the conformational epitope is an amino acid sequence that induces the binding of the antibody to the MUC1\* region on cancer cells. Thus, a term “conformational inducing peptide sequence” may be used, which indicates that a peptide sequence is present within a larger peptide not as a binding site but that induces binding of an antibody to the larger peptide by causing a three-dimensional structure to form that facilitates the binding of the antibody to the larger peptide.

**[00500] MUC1\* antibodies (anti-PSMGFR) for treatment or prevention of cancers**

**[00501]** We discovered that a cleaved form of the MUC1 (SEQ ID NO:1) transmembrane protein is a growth factor receptor that drives the growth of over 75% of all human solid tumor cancers. The cleaved form of MUC1, which we called MUC1\* (pronounced muk 1 star), is a powerful growth factor receptor. Enzymatic cleavage releases the bulk of the MUC1 extracellular domain. It is the remaining portion comprising a truncated extracellular domain, transmembrane domain and cytoplasmic tail that is called MUC1\*. Cleavage and release of the bulk of the extracellular domain of MUC1 unmasks a binding site for activating ligands dimeric NME1, NME6, NME8, NME7<sub>AB</sub>, NME7-X1 or NME7. Cell growth assays show that it is ligand-induced dimerization of the MUC1\* extracellular domain that promotes growth (**Fig. 1A-1D**). MUC1\* positive cells treated with either bivalent ‘bv’ anti-MUC1\* antibody, monovalent ‘mv’ or Fab, NM23-H1 dimers or NME7-AB. Bivalent anti-MUC1\* antibodies stimulate growth of cancer cells whereas the monovalent Fab inhibits growth. Classic bell-shaped curve indicates ligand induced dimerization stimulates growth. Dimeric NM23-H1, aka NME1, stimulates growth of MUC1\* positive cancer cells but siRNA to suppress MUC1 expression eliminate its effect (**Fig. 1C**). NME7-AB also stimulates the growth of MUC1\* positive cells (**Fig. 1D**).

**[00502]** MUC1\* is an excellent target for cancer drugs as it is aberrantly expressed on over 75% of all cancers and is likely overexpressed on an even higher percentage of metastatic

cancers. After MUC1 cleavage, most of its extracellular domain is shed from the cell surface. The remaining portion has a truncated extracellular domain that at least comprises the primary growth factor receptor sequence, PSMGFR (SEQ ID NO:2). Antibodies that bind to the PSMGFR sequence and especially those that competitively inhibit the binding of activating ligands such as NME proteins, including NME1, NME6, NME8, NME7<sub>AB</sub>, NME7-X1 and NME7, are ideal therapeutics and can be used to treat or prevent MUC1 positive or MUC1\* positive cancers, as stand-alone antibodies, antibody fragments or variable region fragments thereof incorporated into bispecific antibodies, or chimeric antigen receptors also called CARs, which are then transfected or transduced into immune cells, then administered to a patient.

**[00503]** Therapeutic anti-MUC1\* antibodies can be monoclonal, polyclonal, antibody mimics, engineered antibody-like molecules, full antibodies or antibody fragments. Examples of antibody fragments include but are not limited to Fabs, scFv, and scFv-Fc. Human or humanized antibodies are preferred for use in the treatment or prevention of cancers. In any of these antibody-like molecules, mutations can be introduced to prevent or minimize dimer formation. Anti-MUC1\* antibodies that are monovalent or bispecific are preferred because MUC1\* function is activated by ligand induced dimerization. Typical binding assays show that NME1 and NME7<sub>AB</sub> bind to the PSMGFR peptide portion of MUC1\* (**Fig. 2A, 2D**). Further, they show that these activating growth factors bind to the membrane proximal portion of MUC1\*, as they do not bind to the PSMGFR peptide if the 10 C-terminal amino acids are missing. Similarly, anti-MUC1\* antibodies MN-C2 and MN-E6 bind to the PSMGFR peptide if and only if the 10 C-terminal amino acids are present (**Fig. 2B, 2C**). Antibodies MN-C3 and MN-C8 bind to epitopes that are different from MN-C2 and MN-E6, as they do not depend on the presence of the 10 C-terminal amino acids of the PSMGFR peptide (**Fig. 2E, 2F**). Antibodies MN-C2, MN-E6, or fragments derived from them, can be administered to a patient for the treatment or prevention of cancers, as stand-alone antibodies or incorporated into bispecific antibodies, BiTEs or chimeric antigen receptors also called CARs that have been transduced into immune cells. MNC2 and MNE6 and other anti-MUC1\* antibodies that competitively inhibit the binding of NME1 and NME7<sub>AB</sub> are preferred for use as stand alone antibody therapeutics.

**[00504]** Therapeutic anti-MUC1\* antibodies for use as a stand alone antibody therapeutic or for integration into a BiTE or a CAR can be selected based on specific criteria. The parent antibody can be generated using typical methods for generating monoclonal antibodies in

animals. Alternatively, they can be selected by screening antibody and antibody fragment libraries for their ability to bind to a MUC1\* peptide, which can be:

**[00505]** (i) PSMGFR region of MUC1;

**[00506]** (ii) PSMGFR peptide;

**[00507]** (iii) a peptide having amino acid sequence of QFNQYKTEAASRYNLTISDVSVDVPFPFSAQSGA (N-10)

**[00508]** (iv) a peptide having amino acid sequence of

**[00509]** ASRYNLTISDVSVDVPFPFSAQSGA (N-19)

**[00510]** (v) a peptide having amino acid sequence of

**[00511]** NLTISDVSVDVPFPFSAQSGA (N-23)

**[00512]** (vi) a peptide having amino acid sequence of

**[00513]** ISDVSVDVPFPFSAQSGA (N-26)

**[00514]** (vii) a peptide having amino acid sequence of

**[00515]** SVSDVPFPFSAQSGA (N-30)

**[00516]** (viii) a peptide having amino acid sequence of

**[00517]** QFNQYKTEAASRYNLTISDVSVDVPFPFS (N-10/C-5)

**[00518]** (ix) a peptide having amino acid sequence of

**[00519]** ASRYNLTISDVSVDVPFPFS (N-19/C-5) or

**[00520]** (x) a peptide having amino acid sequence of

**[00521]** FPFSAQSGA (N-36).

**[00522]** Resultant antibodies or antibody fragments generated or selected in this way can then be further selected by passing additional screens. For example, antibodies or antibody fragments become more preferred based on their ability to bind to MUC1\* positive cancer cells or tissues but not to MUC1 negative cancer cells or to normal tissues. Further, anti-MUC1\* antibodies or antibody fragments may be de-selected as anti-cancer therapeutics if they bind to stem or progenitor cells. Anti-MUC1\* antibodies or antibody fragments become more preferred if they have the ability to competitively inhibit the binding of activating ligands to MUC1\*. **Figs. 3A-3C** shows that MN-E6 and MN-C2 competitively inhibit the binding of activating ligands NME1 and NME7 to MUC1\*.

**[00523]** A process for selecting anti-MUC1\* antibodies for use in treating a patient diagnosed with a MUC1 positive cancer, at risk of developing a MUC1 positive cancer or suspected of

having a MUC1 positive cancer comprises one or more of the following steps of selecting antibodies or antibody fragments that 1) bind to the PSMGFR peptide; 2) bind to the N-10 PSMGFR peptide; 3) bind to cancer cells; 4) do not bind to stem or progenitor cells; and 5) competitively inhibited the binding of dimeric NME1 or NME7-AB to the PSMGFR peptide. For example, **Figs. 3A-3C** show that monoclonals MN-E6 and MN-C2 satisfy all five criteria, while monoclonals MN-C3 and MN-C8 do not competitively inhibit the binding of activating ligands NME1 and NME7 (**Fig. 3C**). Recall that the MUC1\* growth factor receptor is activated by ligand-induced dimerization of its extracellular domain. Therefore, the ideal antibody therapeutic should not dimerize the MUC1\* extracellular domain. Preferably, suitable antibodies in this regard include monovalent antibodies such as those generated in llamas and camels, Fabs, scFv's, single domain antibodies (sdAb), scFv-Fc as long as the Fc portion is constructed such that it does not homo-dimerize.

**[00524]** FACS scans show that anti-MUC1\* antibodies MN-C2 and MN-E6 specifically bind to MUC1\* positive solid tumor cancer cells and MUC1\* transfected cells but not MUC1\* negative or MUC1 negative cells. In one example, a humanized MN-C2 scFv is shown to bind to ZR-75-1, aka 1500, MUC1\* positive breast cancer cells (**Fig. 4A-4C**). MN-E6 was shown to bind to MUC1 negative HCT-116 colon cancer cells if and only if they were transfected with MUC1\*. MN-E6 also bound to MUC1\* positive cancer cells such as ZR-75-1, aka 1500, MUC1\* positive breast cancer cells (**Fig. 4D-4F**). Binding assays such as ELISAs, immunofluorescence, and the like all confirm that MN-C2 and MN-E6 bind to the PSMGFR peptide and to live MUC1 positive cancer cells. Humanized anti-MUC1\* antibodies are selected based on their ability to also bind to the PSMGFR peptide or to MUC1 positive cancer cells. Figure 5 shows that humanized MN-C2 scFv binds with high affinity to the MUC1\* peptide PSMGFR with an EC-50 of about 333nM. Humanized MN-C2 scFv, like Fabs, potently inhibits the growth of MUC1\* positive cancer cells as is shown in one example in **Figs. 6A, 6B**. Like the parent antibodies, humanized scFv's show the same binding pattern. huMNE6-scFv binds to the PSMGFR peptide, binds to the N-10 peptide but does not bind to the C-10 peptide (SEQ ID NO:825) (**Fig. 8**). Murine or humanized MNC3-scFv binds to the, PSMGFR peptide, binds to the N-10 peptide and binds to the C-10 peptide (**Fig. 9**).

**[00525]** The Fabs of MN-E6 and MN-C2 or the comparable single chain variable regions derived from them potently inhibit the growth of MUC1\* positive cancers *in vitro* and *in vivo*. In

several examples, the Fabs of Anti-MUC1\* antibodies inhibited the growth of human MUC1\* positive cancers in vivo. In one case, immune-compromised mice were implanted with human breast tumors then treated with MN-E6 Fab after tumor engraftment. **Fig. 7A** shows that MN-E6 Fab potently inhibited the growth of MUC1\* positive breast cancers. Female nu/nu mice implanted with 90-day estrogen pellets were implanted with 6 million T47D human breast cancer cells that had been mixed 50/50 with Matrigel. Mice bearing tumors that were at least 150 mm<sup>3</sup> and had three successive increases in tumor volume were selected for treatment. Animals were injected sub-cutaneously twice per week with 80 mg/kg MN-E6 Fab and an equal number of mice fitting the same selection criteria were injected with vehicle alone (**Fig. 7A**).

**[00526]** In another aspect, MN-E6 was shown to halt the growth of prostate cancer. **Fig. 7B** shows that MN-E6 Fab potently inhibited the growth of MUC1\* positive prostate cancers. Male NOD/SCID mice were implanted with 6 million DU-145 human prostate cancer cells that had been mixed 50/50 with Matrigel. Mice bearing tumors that were at least 150 mm<sup>3</sup> and had three successive increases in tumor volume were selected for treatment. Animals were injected sub-cutaneously every 48 hours with 160 mg/kg MN-E6 Fab and an equal number of mice fitting the same selection criteria were injected with vehicle alone (**Fig. 7B**). Tumors were measured independently by two researchers twice per week and recorded. Statistics were blindly calculated by independent statistician, giving a P value of 0.0001 for each. Anti-MUC1\* Fab inhibited breast cancer growth and prostate cancer growth. Treatment had no effect on weight, bone marrow cell type or number. The MN-E6 Fab effectively inhibited the growth of the tumors, while the control group's tumors continued to grow until sacrifice. No adverse effects of treatment were observed or detected.

**[00527]** Recombinant forms of MN-E6 and MNC2 were constructed that like the Fab are monomeric. In this case, MN-E6 was humanized and MN-C2 was humanized. There are a number of methods known to those skilled in the art for humanizing antibodies. In addition to humanizing, libraries of human antibodies can be screened to identify other fully human antibodies that bind to the PSMGFR.

**[00528]** A single chain of the humanized MN-E6 variable region, called an scFv, was genetically engineered such that it was connected to the Fc portion of the antibody (SEQ ID NO:256 and 257). Fc regions impart certain benefits to antibody fragments for use as therapeutics. The Fc portion of an antibody recruits complement, which in general means it can



recruit other aspects of the immune system and thus amplify the anti-tumor response beyond just inhibiting the target. The addition of the Fc portion also increases the half-life of the antibody fragment (Czajkowsky DM, Hu J, Shao Z and Pleass RJ. (2012) Fc-fusion proteins: new developments and future perspectives. *EMBO Mol Med.* 4(10):1015-1028). However, the Fc portion of an antibody homo-dimerizes, which in the case of anti-MUC1\* antibody based therapeutics is not optimal since ligand-induced dimerization of the MUC1\* receptor stimulates growth. Therefore, mutations in the Fc region that resist dimer formation are preferred for anti-MUC1\* anti-cancer therapeutics. Deletion of the hinge region and other mutations in the Fc region that make the Fc-mutant resistant to dimerization were made and could be used as therapeutics.

**[00529]** A human or humanized MN-E6 antibody or antibody fragment, Fab, MN-E6 scFv or hu MN-E6 scFv-Fc<sub>mut</sub> are effective anti-cancer agents that can be administered to a person diagnosed with a MUC1 or MUC1\* positive cancer, suspected of having a MUC1 or MUC1\* positive cancer or is at risk of developing a MUC1 or MUC1\* positive cancer.

**[00530] Humanizing**

**[00531]** Humanized antibodies or antibody fragments or fully human antibodies that bind to the extracellular domain of -MUC1\* are preferred for therapeutic use. The techniques described herein for humanizing antibodies are but a few of a variety of methods known to those skilled in the art. The invention is not meant to be limited by the technique used to humanize the antibody.

**[00532]** Humanization is the process of replacing the non-human regions of a therapeutic antibody (usually mouse monoclonal antibody) by human one without changing its binding specificity and affinity. The main goal of humanization is to reduce immunogenicity of the therapeutic monoclonal antibody when administered to human. Three distinct types of humanization are possible. First, a chimeric antibody is made by replacing the non-human constant region of the antibody by the human constant region. Such antibody will contain the mouse Fab region and will contain about 80-90% of human sequence. Second, a humanized antibody is made by grafting of the mouse CDR regions (responsible of the binding specificity) onto the variable region of a human antibody, replacing the human CDR (CDR-grafting method). Such antibody will contain about 90-95% of human sequence. Third and last, a full human antibody (100% human sequence) can be created by phage display, where a library of human

antibodies is screened to select antigen specific human antibody or by immunizing transgenic mice expressing human antibody.

**[00533]** A general technique for humanizing an antibody is practiced approximately as follows. Monoclonal antibodies are generated in a host animal, typically in mice. Monoclonal antibodies are then screened for affinity and specificity of binding to the target. Once a monoclonal antibody that has the desired effect and desired characteristics is identified, it is sequenced. The sequence of the animal-generated antibody is then aligned with the sequences of many human antibodies in order to find human antibodies with sequences that are the most homologous to the animal antibody. Biochemistry techniques are employed to paste together the human antibody sequences and the animal antibody sequences. Typically, the non-human CDRs are grafted into the human antibodies that have the highest homology to the non-human antibody. This process can generate many candidate humanized antibodies that need to be tested to identify which antibody or antibodies has the desired affinity and specificity.

**[00534]** Once a human antibody or a humanized antibody has been generated it can be further modified for use as an Fab fragment, as a full antibody, or as an antibody-like entity such as a single chain molecule containing the variable regions, such as scFv or an scFv-Fc. In some cases it is desirable to have Fc region of the antibody or antibody-like molecule mutated such that it does not dimerize.

**[00535]** In addition to methods that introduce human sequences into antibodies generated in non-human species, fully human antibodies can be obtained by screening human antibody libraries with a peptide fragment of an antigen. A fully human antibody that functions like MN-E6 or MN-C2 is generated by screening a human antibody library with a peptide having the sequence of the PSMGFR N-10 peptide. Humanized anti-MUC1\* antibodies were generated based on the sequences of the mouse monoclonal antibodies MN-E6 and MN-C2. In one aspect of the invention, a patient diagnosed with a MUC1\* positive cancer is treated with an effective amount of a murine or camelid MNC2, MNE6, 20A10 (SEQ ID NOS:1574-1581), 3C2B1 (SEQ ID NOS:1572-1573), 5C6F3, 25E6 (SEQ ID NO:1598-1601), 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11. In another aspect of the invention, a patient diagnosed with a MUC1\* positive cancer is treated with an effective amount of humanized MN-E6 or MN-C2. In a preferred embodiment, a patient diagnosed with a MUC1\* positive cancer is treated with an effective amount of humanized MNC2 , MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9,

1E4, B12, B2, B7, B9, 8C7F3, or H11. In another aspect of the invention, a patient diagnosed with a MUC1\* positive cancer is treated with an effective amount of humanized monovalent MNC2, MNE6, 20A10 (SEQ ID NOS:1574-1581), 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11, wherein monovalent means the corresponding Fab fragment, the corresponding scFv or the corresponding scFv-Fc fusion. In a preferred embodiment, a patient diagnosed with a MUC1\* positive cancer is treated with an effective amount of a humanized scFv or monomeric humanized scFv-Fc of MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11. Since the MUC1\* growth factor receptor is activated by ligand induced dimerization of its extracellular domain, and because the Fc portion of an antibody homo-dimerizes, it is preferable that a construct that includes an Fc portion uses a mutated Fc region that prevents or minimizes dimerization.

**[00536]** Antibodies that bind to PSMGFR (SEQ ID NO:2) peptide, and more specifically to the N-10 peptide, of the extracellular domain of the MUC1\* receptor are potent anti-cancer therapeutics that are effective for the treatment or prevention of MUC1\* positive cancers. They have been shown to inhibit the binding of activating ligands dimeric NME1 (SEQ ID NO:1781) and NME7<sub>AB</sub> (SEQ ID NOS:827) to the extracellular domain of MUC1\*. Anti-MUC1\* antibodies that bind to the PSMGFR sequence inhibit the growth of MUC1\*-positive cancer cells, specifically if they inhibit ligand-induced receptor dimerization. Fabs of anti-MUC1\* antibodies have been demonstrated to block tumor growth in animals. Thus, antibodies or antibody fragments that bind to the extracellular domain of MUC1\* would be beneficial for the treatment of cancers wherein the cancerous tissues express MUC1\*.

**[00537]** Antibodies that bind to PSMGFR region of MUC1\* or bind to a synthetic PSMGFR peptide are preferred. We have identified several monoclonal antibodies that bind to the extracellular domain of MUC1\*. Among this group are mouse monoclonal antibodies MNC2 (SEQ ID NOS:118-131, 144-158, 163-164, 168-181, 194-209), MNE6 (SEQ ID NOS:12-25, 39-59, 65-78, 93-114), 20A10 (SEQ ID NOS:988-1019, 1574-1597, 1659-1666); 3C2B1 (SEQ ID NOS:1386-1413, 1572-1573), 5C6F3 (SEQ ID NOS:1356-1385), 25E6 (SEQ ID NOS:1020-1051, 1598-1617, 1667-1674), 18G12 (SEQ ID NOS:956-987), 28F9 (SEQ ID NOS:1052-1083), 1E4 (SEQ ID NOS:1116-1227), B12 (SEQ ID NOS:1414-1431, 1733-1742), B2 (SEQ ID NOS:1432-1459), B7 (SEQ ID NOS:1460-1487), B9 (SEQ ID NOS:1544-1571), 8C7F3 (SEQ ID NOS:1488-1515), or H11 (SEQ ID NOS:1516-1543), the variable regions of which were

sequenced and are given as for MN-E6 SEQ ID NOS: 12-13 and 65-66, for MN-C2 SEQ ID NOS: 118-119 and 168-169. The CDRs of these antibodies make up the recognition units of the antibodies and are the most important parts of the mouse antibody that should be retained when grafting into a human antibody. The sequences of the CDRs for each mouse monoclonal are as follows, heavy chain sequence followed by light chain: MN-E6 CDR1 (SEQ ID NO:16-17 and 69-70) CDR2 (SEQ ID NO:20-21 and 73-74) CDR3 (SEQ ID NO: 24-25 and 77-78), MN-C2 CDR1 (SEQ ID NO:122-123 and 172-173) CDR2 (SEQ ID NO:126-127 and 176-177) CDR3 (SEQ ID NO:130-131 and 180-181). In some cases, portions of the framework regions that by modeling are thought to be important for the 3-dimensional structure of the CDRs, are also imported from the mouse sequence.

**[00538]** Monoclonal antibodies MN-E6 and MN-C2 have greater affinity for MUC1\* as it appears on cancer cells. Monoclonal antibodies MN-C3 and MN-C8 have greater affinity for MUC1\* as it appears on stem cells.

**[00539]** All four antibodies have been humanized, which process has resulted in several humanized forms of each antibody. CDRs derived from the variable regions of the mouse antibodies were biochemically grafted into a homologous human antibody variable region sequence. Humanized variable regions of MN-E6 (SEQ ID NOS: 38-39 and 93-94), MN-C2 (SEQ ID NOS: 144-145 and 194-195), MN-C3 (SEQ ID NOS: 439-440 and 486-487) and MN-C8 (SEQ ID NOS: 525-526 and 543-544) were generated by grafting the mouse CDRs into the variable region of a homologous human antibody. The humanized heavy chain variable constructs were then fused into constant regions of either human IgG1 heavy chain constant region (SEQ ID NOS:58-59) or human IgG2 heavy chain constant region (SEQ ID NO:54-55), which are then paired with either humanized light chain variable constructs fused to a human kappa chain (SEQ ID NO: 109-110) or human lambda chain (SEQ ID NO: 113-114) constant region. Other IgG isotypes could be used as constant region including IgG3 or IgG4.

**[00540]** Examples of humanized MN-E6 variable region into an IgG2 heavy chain (SEQ ID NOS:52-53) and into an IgG1 heavy chain (SEQ ID NOS:56-57), humanized MN-C2 variable into an IgG1 heavy chain (SEQ ID NOS: 157-158) or into an IgG2 heavy chain (SEQ ID NOS: 163-164) paired with either Lambda light chain (SEQ ID NO: 111-112 and 216-219) or Kappa chain (SEQ ID NO:107-108 and 210-213) and , humanized MN-C3 (SEQ ID NOS: 455-456, 453-454 and 500-501, 502-503) and MN-C8 (SEQ ID NOS: 541-542, 539-540 and 579-580,

581-582) antibodies were generated. Which IgG constant region is fused to the humanized variable region depends on the desired effect since each isotype has its own characteristic activity. The isotype of the human constant region is selected on the basis of things such as whether antibody dependent cell cytotoxicity (ADCC) or complement dependent cytotoxicity (CDC) is desired but can also depend on the yield of antibody that is generated in cell-based protein expression systems. In a preferred embodiment, humanized anti-MUC1\* antibodies or antibody fragments are administered to a person diagnosed with or at risk of developing a MUC1-positive cancer.

**[00541]** One method for testing and selecting the humanized anti-MUC1\* antibodies that would be most useful for the treatment of persons with cancer or at risk of developing cancers is to test them for their ability to inhibit the binding of activating ligands to the MUC1\* extracellular domain. Dimeric NME1 can bind to and dimerize the MUC1\* extracellular domain and in so doing stimulates cancer cell growth. Antibodies and antibody fragments that compete with NME1 for binding to the MUC1\* extracellular domain are therefore anti-cancer agents. NME7<sub>AB</sub> is another activating ligand of MUC1\*. In some cases, it is preferable to identify antibodies that block the binding of NME7, or an NME7<sub>AB</sub> truncation or cleavage product of NME7-X1, to the MUC1\* extracellular domain. Antibodies and antibody fragments that compete with NME7 and NME7 variants for binding to the MUC1\* extracellular domain are effective as anti-cancer therapeutics. These antibodies include but are not limited to MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11 as well as single chain versions, such as scFv, of these antibodies and humanized version thereof. Other NME proteins also bind to MUC1 or MUC1\* including NME6 and NME8. Antibodies that compete with these proteins for binding to MUC1\* may also be useful as therapeutics. In a preferred embodiment, murine, camelid, human or humanized anti-MUC1\* antibodies or antibody fragments are administered to a person diagnosed with or at risk of developing a MUC1-positive cancer. In a more preferred embodiment, single chain antibody fragments, or monomeric scFv-Fc fusions, derived from humanized sequences of MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11 are administered to a person diagnosed with or at risk of developing a MUC1-positive cancer.

**[00542]** Single chain variable fragments, scFv, or other forms that result in a monovalent antibody or antibody-like protein are also useful. In some cases it is desired to prevent

dimerization of the MUC1\* extracellular domain. Single chain variable fragments, Fabs and other monovalent antibody-like proteins have been shown to be effective in binding to the extracellular domain of MUC1\* and blocking MUC1\* dimerization. These single chain variable fragments, Fabs and other monovalent antibody-like molecules effectively blocked cancer growth *in vitro* and in animals xenografted with human MUC1-positive cancer cells. Thus, humanized single chain variable fragments or monovalent anti-MUC1\* antibodies or antibody-like molecules would be very effective as an anti-cancer therapeutic. Such humanized single chain antibodies, Fabs and other monovalent antibody-like molecules that bind to the MUC1\* extracellular domain or to a PSMGFR peptide are therefore useful as anti-cancer therapeutics. Anti-MUC1\* single chain variable fragments are generated by grafting non-human CDRs of antibodies, which bind to extracellular domain of MUC1\* or bind to PSMGFR peptide, into a framework of a homologous variable region human antibody. The resultant humanized heavy and light chain variable regions are then connected to each other via a suitable linker, wherein the linker should be flexible and of length that it allows heavy chain binding to light chain but discourages heavy chain of one molecule binding to the light chain of another. For example a linker of about 10-15 residues. Preferably, the linker includes [(Glycine)<sub>4</sub> (Serine)<sub>1</sub>]<sub>3</sub> (SEQ ID NOS: 401-402), but is not limited to this sequence as other sequences are possible.

**[00543]** In one aspect, the humanized variable regions of MN-E6 (SEQ ID NOS: 38-39 and 93-94), MN-C2 (SEQ ID NOS: 144-145 and 194-195), or other antibodies of the invention are biochemically grafted into a construct that connects heavy and light chains via a linker. Examples of humanized single chain anti-MUC1\* antibodies comprising humanized sequences from the variable regions of MN-E6 and MN-C2, were generated. Several humanized MN-E6 single chain proteins were generated (SEQ ID NOS: 232-237). Several humanized MN-C2 single chain proteins were generated (SEQ ID NOS: 238-243). In a preferred embodiment, humanized anti-MUC1\* antibody fragments, including variable fragments, scFv antibody fragments MN-E6 scFv, MN-C2 scFv, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11 scFv are administered to a person diagnosed with or at risk of developing a MUC1-positive cancer.

**[00544]** One aspect of the invention is a method for treating a patient diagnosed with, suspected of having, or at risk of developing a MUC1 positive or MUC1\* positive cancer, wherein the patient is administered an effective amount of a monomeric MN-E6 scFv, MN-C2

scFv, or MN-E6 scFv-Fc, MN-C2 scFv-Fc, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11, wherein the antibody variable fragment portions are human or have been humanized and wherein the Fc portion of the antibody-like protein has been mutated such that it resists dimer formation.

**[00545] CAR T and cancer immunotherapy techniques**

**[00546]** In another aspect of the invention, some or all of the single chain portions of anti-MUC1\* antibody fragments are biochemically fused onto immune system molecules, using several different chimeric antigen receptor, 'CAR' strategies. The idea is to fuse the recognition portion of an antibody, typically as a single chain variable fragment, to an immune system molecule that has a transmembrane domain and a cytoplasmic tail that is able to transmit signals that activate the immune system. The recognition unit can be an antibody fragment, a single chain variable fragment, scFv, or a peptide. In one aspect, the recognition portion of the extracellular domain of the CAR is comprised of sequences from the humanized variable region of MN-E6 (SEQ ID NOS:38-39 and 93-94), MN-C2 (SEQ ID NOS:144-145 and 194-195), 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11. Examples of murine or humanized antibodies of the invention, or their single chain fragments, scFv's, which can be incorporated into CARs, BiTEs or ADCs are given as: 3C2B1 (SEQ ID NOS: 1572-1573), 20A10 (SEQ ID NOS: 1574-1581), 25E6 (SEQ ID NOS:1598-1601). In another aspect, it is comprised of sequences from a single chain variable fragment. Examples of single chain constructs are given. Several humanized MN-E6 single chain proteins, scFv, were generated (SEQ ID NOS: 232-237). Several humanized MN-C2 single chain proteins, scFv, were generated (SEQ ID NOS: 238-243). The transmembrane region of the CAR can be derived from CD8, CD4, antibody domains or other transmembrane region, including the transmembrane region of the proximal cytoplasmic co-stimulatory domain, such as CD28, 4-1BB or other. The cytoplasmic tail of the CAR can be comprised of one or more motifs that signal immune system activation. This group of cytoplasmic signaling motifs, sometimes referred to as, co-stimulatory cytoplasmic domains, includes but is not limited to CD3-zeta, CD27, CD28, 4-1BB, OX40, CD30, CD40, ICAM-1, LFA-1, ICOS, CD2, CD5, CD7 and Fc receptor gamma domain. A minimal CAR may have the CD3-zeta or an Fc receptor gamma domain then one or two of the above domains in tandem on the cytoplasmic tail. In one aspect, the cytoplasmic tail comprises CD3-zeta, CD28, 4-1BB and/or OX40.

**[00547]** The extracellular domain recognition unit of a MUC1\* targeting CAR can comprise variable regions of any non-human, humanized or human antibody that is able to bind to at least 12 contiguous amino acids of the PSMGFR peptide (SEQ ID NO:2) or the N-10 peptide. In one aspect, the MUC1\* targeting portion of the CAR comprises variable regions from non-human, humanized or human MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11. Examples of a few antibodies of the invention, incorporated into CARs as either murine or humanized are given as 20A10 (SEQ ID NOS:1582-1597) and 25E6 (SEQ ID NOS:1602-1617). In the humanization process, the antibody CDRs can be inserted into a number of different framework regions; as a demonstration we generated three versions of a humanized 20A10 which differ only in the framework regions. These have been incorporated into CARs (SEQ ID NOS:1675, 1678, 1685) that when transduced into human T cells are able to recognize target MUC1\* expressing cells and kill them. In one aspect, the extracellular domain recognition unit of a CAR is comprised essentially of a humanized MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11 single chain variable fragment scFv. The transmembrane region of the CAR can be derived from CD8 (SEQ ID NOS:363-364), or can be the transmembrane domain of CD3-zeta, CD28, 41bb, OX40 or other transmembrane region (SEQ ID NOS:361-372) and the cytoplasmic domain of a CAR with antibody fragment targeting MUC1\* extracellular domain can be comprised of one or more selected from the group comprising an immune system co-stimulatory cytoplasmic domain. The group of immune system co-stimulatory domains includes but is not limited to CD3-zeta, CD27, CD28, 4-1BB, OX40, CD30, CD40, ICAM-1, LFA-1, ICOS, CD2, CD5, CD7 and Fc receptor gamma domain (SEQ ID NOS:373-382).

**[00548]** The CARs described can be transfected or transduced into a cell of the immune system. In a preferred embodiment, a MUC1\* targeting CAR is transfected or transduced into a T cell. In one aspect, the T cell is a CD3+/CD28+ T cell. In another case it is a dendritic cell. In another case it is a B cell. In another case it is a mast cell. In yet another case it is a Natural Killer, NK, cell. The recipient cell can be from a patient or from a donor. If from a donor, it can be engineered to remove molecules that would trigger rejection. Cells transfected or transduced with a CAR of the invention can be expanded *ex vivo* or *in vitro* then administered to a patient. Administrative routes are chosen from a group containing but not limited to bone marrow transplant, intravenous injection, *in situ* injection or transplant. In a preferred embodiment, the



MUC1\* targeting CAR is administered to a person diagnosed with or at risk of developing a MUC1-positive cancer.

**[00549]** There are many possible anti-MUC1\* CAR constructs that can be transduced into T cells or other immune cells for the treatment or prevention of MUC1\* positive cancers. CARs are made up of modules and the identity of some of the modules is relatively unimportant, while the identity of other modules is critically important.

**[00550]** We and others have shown that intracellular signaling modules, such as CD3-zeta (SEQ ID NOS: 373-376), CD28 (SEQ ID NOS: 377-378) and 41BB (SEQ ID NOS: 379-380), alone or in combinations stimulate immune cell expansion, cytokine secretion and immune cell mediated killing of the targeted tumor cells (Pulè MA, Straathof KC, Dotti G, Heslop HE, Rooney CM and Brenner MK (2005) A chimeric T cell antigen receptor that augments cytokine release and supports clonal expansion of primary human T cells. *Mol Ther.* 12(5):933-941; Hombach AA, Heiders J, Foppe M, Chmielewski M and Abken H. (2012) OX40 costimulation by a chimeric antigen receptor abrogates CD28 and IL-2 induced IL-10 secretion by redirected CD4(+) T cells. *Oncoimmunology.* 1(4):458-466; Kowolik CM, Topp MS, Gonzalez S, Pfeiffer T, Olivares S, Gonzalez N, Smith DD, Forman SJ, Jensen MC and Cooper LJ. (2006) CD28 costimulation provided through a CD19-specific chimeric antigen receptor enhances in vivo persistence and antitumor efficacy of adoptively transferred T cells. *Cancer Res.* 66(22):10995-11004; Loskog A, Giandomenico V, Rossig C, Pule M, Dotti G and Brenner MK. (2006) Addition of the CD28 signaling domain to chimeric T-cell receptors enhances chimeric T-cell resistance to T regulatory cells. *Leukemia.* 20(10):1819-1828; Milone MC, Fish JD, Carpenito C, Carroll RG, Binder GK, Teachey D, Samanta M, Lakhai M, Gloss B, Danet-Desnoyers G, Campana D, Riley JL, Grupp SA and June CH. (2009) Chimeric receptors containing CD137 signal transduction domains mediate enhanced survival of T cells and increased antileukemic efficacy in vivo. *Mol Ther.* 17(8):1453-1464; Song DG, Ye Q, Carpenito C, Poussin M, Wang LP, Ji C, Figini M, June CH, Coukos G, Powell DJ Jr. (2011) In vivo persistence, tumor localization, and antitumor activity of CAR-engineered T cells is enhanced by costimulatory signaling through CD137 (4-1BB). *Cancer Res.* 71(13):4617-4627). Antibodies of the invention including but not limited to fragments of MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11 can also be incorporated into CARs that have mutated cytoplasmic tails, such as mutated tyrosines or ITAMs. In any of the CARs described

above, the cytoplasmic tails may include mutations that dampen signaling. Such mutations include but are not limited to Tyrosines that are mutated to inhibit phosphorylation and signaling (Salter et al, 2018; ). In any of the CARs described above, the ITAMs of CD3-zeta may be mutated to inhibit or dampen signaling (Feucht et al 2019). In any of the CARs described above, the CD3 of the cytoplasmic tail may comprise mutations in the ITAMs including those referred to as 1XX. Examples of antibodies of the invention incorporated into CARs with 1XX mutations in ITAMs of CD3-zeta are given in the following sequences: MNC2 (SEQ ID NOS: 1618-1625), MNE6 (SEQ ID NOS:1626-1633), 20A10 (SEQ ID NOS:1590-1595), 25E6 (SEQ ID NOS:1610-1617). We note that the CDRs of antibodies can be inserted into a background of a number of different framework regions. As an example, 20A10 CDRs were inserted into three different sets of framework regions (SEQ ID NOS:1692, 1699 and 1706) and all were able to function when transduced into T cells. In any of the CARs described above, the T cell may be engineered to overexpress c-Jun as a method to inhibit T cell exhaustion (Lynn et al 2019). A variety of promoters can be used upstream of the genes for CARs and other compositions of the invention, including insertion into a naturally occurring promoter in the cell, such as the TRAC locus, using CRISPR, Sleeping Beauty or similar technology for site directed insertion of a gene. Among the promoters commonly used are the CMV promoter, or a mini CMV (SEQ ID NO: 1634), a minimal IL-2 promoter (SEQ ID NO: 1635), or Minimal Promoter minip (SEQ ID NO: 1636).

**[00551]** Single chain antibody fragments that included the variable domain of the monoclonal anti-MUC1\* antibodies called MN-E6 or MN-C2 were engineered into a panel of CARs. The MUC1\* targeting CARs were then transduced, separately or in combinations, into immune cells. When challenged with surfaces presenting a MUC1\* peptide, an antigen presenting cell transfected with MUC1\*, or MUC1\* positive cancer cells, the immune cells that were transduced with MUC1\* targeting CARs elicited immune responses, including cytokine release, killing of the targeted cells and expansion of the immune cells.

**[00552]** For example, the gene encoding the CARs and activated T cell induced genes described herein can be virally transduced into an immune cell using viruses, or inserted into a region downstream of one of the cell's promoters or enhancers, such as the TRAC (T cell receptor alpha chain) locus. Virus delivery systems and viral vectors including but not limited to retroviruses, including gamma-retroviruses, lentivirus, adenoviruses, adeno-associated viruses,

baculoviruses, poxvirus, herpes simplex viruses, oncolytic viruses, HF10, T-Vec and the like can be used. In addition to viral transduction, CARs and activated T cell induced genes described herein can be directly spliced into the genome of the recipient cell using methods such as CRISPR technology, CRISPR-Cas9 and -CPF1, TALEN, Sleeping Beauty transposon system, and SB 100X.

**[00553]** Similarly, the identity of molecules that make up the non-targeting portions of the CAR such as the extracellular domain, transmembrane domain and membrane proximal portion of the cytoplasmic domain, are not essential to the function of a MUC1\*-targeting CAR. For example, the extracellular domain, transmembrane domain and membrane proximal portion of the cytoplasmic domain can be comprised of portions of CD8, CD4, CD28, or generic antibody domains such as Fc, CH2CH3, or CH3. Further, the non-targeting portions of a CAR can be a composite of portions of one or more of these molecules or other family members.

**[00554]** One aspect of the invention is a method for treating a patient diagnosed with, suspected of having, or at risk of developing a MUC1 positive or MUC1\* positive cancer, wherein the patient is administered an effective amount of immune cells that have been transduced with a MUC1\* targeting CAR. In another aspect of the invention, the immune cells are T cells isolated from a patient, which are then transduced with CARs wherein the targeting head of the CAR binds to MUC1\*, and after expansion of transduced T cells, the CAR T cells are administered in an effective amount to the patient. In yet another aspect of the invention, the immune cells are T cells isolated from a patient, which are then transduced with CARs wherein the targeting head of the CAR comprises portions of MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11, and after optional expansion of transduced T cells, the CAR T cells are administered in an effective amount to the patient.

**[00555] Specificity of anti-MUC1\* targeting antibodies**

**[00556]** As these experiments demonstrate, the critical portion of a CAR is the antibody fragment that directs the immune cell to the tumor cell. As we will show in the following section, MN-E6 and MN-C2 are specific for the form of MUC1\* that is expressed on tumor cells. The next most important part of a CAR is the cytoplasmic tail bearing immune system co-stimulatory domains. The identity of these domains modulates the degree of immune response but does not affect the specificity. As shown, the identity of the transmembrane portion of a CAR is the least important. It appears that as long as the transmembrane portion has some flexibility and is long

enough to allow the antibody fragment to reach its cognate receptor on the tumor cell, it will suffice. CARs comprising the MN-E6 targeting antibody fragment, and intracellular co-stimulatory domains 41BB and CD3-zeta but having a variety of different extracellular, transmembrane and short cytoplasmic tail all worked in that they specifically killed the targeted cells while stimulating the expansion of the host T cells.

**[00557]** The most accurate way of demonstrating antibody specificity is testing the antibody on normal human tissue specimens compared to cancerous tissue specimens. MN-C2 and MN-E6 were shown to specifically bind to MUC1 or MUC1\* positive cancer cells. Several breast tumor arrays were assayed using several anti-MUC1 or MUC1\* antibodies. Essentially the studies involving serial sections of breast cancer tissue specimens from over 1,200 different breast cancer patients showed that very little full-length MUC1 remains on breast cancer tissues. The vast majority of the MUC1 expressed is MUC1\* and is stained by MN-C2. The analysis was performed by Clariant Diagnostics and tissue staining was scored using the Allred method. For example, **Fig. 10** shows serial sections of breast cancer tissue arrays that were stained with either VU4H5, a commercially available anti-MUC1 antibody that binds to the tandem repeats, or MN-C2 that binds to MUC1\*. **Figs. 10** and **11** are photographs of breast cancer tissue arrays stained with either VU4H5 which recognizes MUC1-FL (full length) or MN-C2 which recognizes cancerous MUC1\*. Tissue staining was scored using Allred scoring method which combines an intensity score and a distribution score. Below the photographs of the tissue arrays are color-coded graphs displaying the results. As can be seen, the arrays stained with VU4H5 are very light and many tissues do not stain at all despite the published reports that MUC1 is aberrantly expressed on over 96% of all breast cancers as evidenced by nucleic acid based diagnostics. In contrast, the arrays stained with MN-C2 are very dark (red versus yellow or white in graph). Additionally, many tissues did not stain at all with anti-full-length MUC1 but stained very dark with MN-C2, (see green boxes in graph). Similarly, we stained normal or cancerous breast tissues with humanized MN-E6 scFv-Fc. The antibody fragment was biotinylated so it could be visualized by a secondary streptavidin based secondary. As can be seen in **Fig. 12**, hMN-E6 scFv-Fc does not stain normal breast tissue but stains cancerous breast tissue. Further, the intensity and homogeneity of staining increases with tumor grade and/or metastatic grade of the patient (**Fig. 12-13**). Similarly, hMN-E6 scFv-Fc did not stain normal lung tissue but did stain lung cancer tissue (**Fig. 14-18**) and the intensity and distribution of staining increased as tumor

grade or metastatic grade increased. **Fig. 19** shows photographs of normal small intestine and cancerous small intestine tissues stained with humanized MN-E6-scFv-Fc biotinylated anti-MUC1\* antibody at 5 ug/mL, then stained with a secondary streptavidin HRP antibody. A) is a normal small intestine tissue. B) is small intestine cancer from patient as denoted in the figure. C,D are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 20** shows photographs of normal small intestine tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are normal small intestine tissue. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 21** shows photographs of cancerous small intestine tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are cancerous small intestine tissue from a patient as denoted in figure. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 22** shows photographs of cancerous small intestine tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are cancerous small intestine tissue from a patient as denoted in figure. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 23** shows photographs of normal colon tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are normal colon. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 24** shows photographs of colon cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are colon cancer tissue from a metastatic patient as denoted in figure. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 25** shows photographs of colon cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are colon cancer tissue from a Grade 2 patient as denoted in figure. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 26** shows photographs of colon cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are

colon cancer tissue from a metastatic patient as denoted in figure. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 27** shows photographs of prostate cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are prostate cancer tissue from a patient as denoted in figure. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 28** shows photographs of prostate cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are prostate cancer tissue from a patient as denoted in figure. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone. **Fig. 29** shows photographs of prostate cancer tissues stained with humanized MN-E6-scFv-Fc anti-MUC1\* antibody at 50 ug/mL, then stained with a secondary goat-anti-human HRP antibody. A-D are prostate cancer tissue from a patient as denoted in figure. E-H are photographs of the corresponding serial sections that were stained with the secondary antibody alone.

**[00558]** One aspect of the invention is a method for treating a patient diagnosed with, suspected of having, or at risk of developing a MUC1 positive or MUC1\* positive cancer, wherein a specimen is obtained from the patient's cancer and is tested for reactivity with an antibody that binds to PSMGFR SEQ ID NO:2, or more specifically to the N-10 peptide. The patient is then treated with an scFv, scFv-Fc or CAR T that comprises antibody variable fragments from the antibody that reacted with their cancer specimen or can be chosen from among MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11. Another aspect of the invention is a method for treating a patient diagnosed with, suspected of having, or at risk of developing a MUC1 positive or MUC1\* positive cancer, wherein a specimen is obtained from the patient's cancer and is tested for reactivity with MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11; the patient is then treated with the antibody, antibody fragment, scFv, scFv-Fc-mut, BiTE or CAR T that comprises portions of the antibody that reacted with their cancer specimen.

**[00559]** As we previously reported, it is MUC1\*, the transmembrane cleavage product, not full-length MUC1, the is a growth factor receptor that drives tumor growth. The growth factors that activate MUC1\* bind to ectopic sites that are only exposed after cleavage and release of the tandem repeat portion of MUC1. Antibodies of the invention, like the activating growth factors,

cannot bind to full-length MUC1. FACS analysis clearly shows that anti-MUC1\* antibody MNC2 is unable to bind to HCT-116, MUC1 negative cells (Fig. 35A), binds robustly to those cells if they are transfected with MUC1\* (Fig. 35B), but will not bind to HCT cells transfected with full-length MUC1 (Fig. 35C). A commercially available anti-tandem repeat antibody VU4H5 clearly recognizes full-length MUC1 (Fig. 35D).

**[00560]** We discovered that MUC1 can be cleaved to MUC1\* by more than one cleavage enzyme and that the site of cleavage affects its fold and consequently affects which monoclonal antibody is able to recognize that form of MUC1\*. Different cancer cells or cancerous tissues express different cleavage enzymes. We tested various cleavage enzyme inhibitors on different cancer cell lines and found that an inhibitor that inhibits cleavage of MUC1 in one cancer cell line did not inhibit its cleavage in another cancer cell line. Similarly, PCR experiments showed that cleavage enzymes are expressed at different levels in different cells or cell lines. For example, hematopoietic stem cells of the bone marrow express a MUC1\* that is recognized by monoclonal antibody MNC3 but not MNE6 or MNC2 (**Fig. 39**). The growth of DU145 prostate cancer cells and T47D breast cancer cells is inhibited by the Fabs of MNC2 and MNE6 but not by the Fabs of MNC3 or MNC8, indicating that the cancer cell lines express a MUC1\* that is recognized by MNE6 and MNC2 but not by MNC3 or MNC8 (**Fig. 42**). PCR experiments show that CD34 positive cells of the bone marrow express about 2,500-times more MMP2 and about 350-times more ADAM28 than T47D breast cancer cells, while DU145 prostate cancer cells express about 2,000-times more ADAM TS16, about 400-times more MMP14 and about 100-times more MMP1 than T47D breast cancer cells (**Fig. 43 and Fig. 44**). Conversely, T47D breast cancer cells express about 80-times more MMP9 than the bone marrow cells and about twice as much as DU145 prostate cancer cells. Various cleavage enzyme inhibitors were tested for their ability to inhibit cleavage in different kinds of cancer cells.

**[00561] General strategy for using antibodies, antibody fragments and CARs that target the extracellular domain of MUC1\***

**[00562]** In one aspect of the invention, a second factor, which may be a cleavage enzyme, an antibody, a cytokine, or a second CAR, and a CAR are transduced into the same T cell. In another aspect of the invention, the second factor is on an inducible promoter such that its expression is activated when the CAR engages the targeted cancer cells. In some cases, the expression of the second factor is controlled by an inducible promoter. In one aspect of the

invention, expression of the second factor is induced when the immune cell is activated, for example when it recognizes or engages its target. In one example, a T cell is transfected or transduced with a second factor whose expression is induced when the T cell recognizes a target cancer cell. One way to do this is to induce expression of the second factor when, or shortly after, an NFAT protein is expressed or translocated to the nucleus. For example, a sequence derived from an NFAT promoter region is put upstream of the gene for the second factor. In this way, when the transcription factors that bind to the promoter of the NFAT protein are present in sufficient concentration to bind to and induce transcription of the NFAT protein, they will also bind to that same promoter that is engineered in front of the sequence for transcription of the second factor. The NFAT protein may be NFAT1 also known as NFATc2, NFAT2 also known as NFATc or NFATc1, NFAT3 also known as NFATc4, NFAT4 also known as NFATc3, or NFAT5. In one aspect of the invention, the NFAT is NFATc1, NFATc3 or NFATc2. In one aspect of the invention, the NFAT is NFAT2 also known as NFATc1. SEQ ID NO:646 shows nucleic acid sequence of the upstream transcriptional regulatory region for NFAT2. The promoter sequence for NFAT gene may include the nucleic acid sequence of SEQ ID NO:781-783 or SEQ ID NO:815 as examples, but it can be seen that the optimal sequence or minimal sequence for expression of the second factor may be obtained by making fragments, extensions or mutations of the promoter and testing for the strength of the promoter with respect to expression of the second factor. In one aspect of the invention, the transcriptional regulatory region for NFAT2 is engineered upstream of the gene encoding the second factor, which if for cleavage enzyme MMP9 (SEQ ID NO:647) or the catalytic sub-unit of MMP9 (SEQ ID NO:648). In one aspect of the invention, the NFAT is NFATc3 and the promoter sequence of NFATc3 includes nucleic acid sequences from SEQ ID NO:816. In one aspect of the invention, the transcriptional regulatory region for NFATc3 is engineered upstream of the gene encoding the second factor, here as an example is MMP9. In another aspect of the invention, the NFAT is NFATc2. SEQ ID NO:817-818 shows nucleic acid sequence of the upstream transcriptional regulatory region for NFATc2. In one aspect of the invention, the transcriptional regulatory region for NFATc2 is engineered upstream of the gene encoding the second factor, which may be cleavage enzyme MMP9 (SEQ ID NO:647) or the catalytic sub-unit of MMP9 (SEQ ID NO:648).



**[00563]** Another method for having the expression of the second factor induced when the T cell or CAR T cell is activated is to have the gene for the second factor on an inducible promoter where the NFAT protein itself binds to and induces transcription of the second factor. In this case, an NFAT response element (NFAT RE) may be positioned upstream of the gene for the second factor or fragment of the second factor. The NFAT may bind to its responsive element upstream of the second factor alone or as part of a complex. The NFAT protein may be NFATc1, NFATc2, NFATc3, NFATc4, or NFAT5. In a preferred embodiment, the NFAT protein is NFAT2 aka NFATc1, aka NFATc. The gene of the second factor or fragment thereof is cloned downstream of an NFAT-response element (SEQ ID NO:649), which may be repeats of the response element (SEQ ID NO:650) and CMV minimal promoter (mCMV) (SEQ ID NO:651) to induce expression of second factor by NFAT protein. The NFAT response element may include nucleic acid sequence of NFAT consensus sequence (SEQ ID NO:804). The NFAT response element may include the nucleic acid sequence of SEQ ID NOS:805-814 as examples, but it can be seen that the optimal sequence or minimal sequence for expression of the second factor may be obtained by making fragments, extensions or mutations of the responsive element nucleic acid and testing for the strength of the responsive element with respect to expression of the second factor. The enhancer region of Foxp3 also contains NFAT response elements within the 120-bp from 2079 to 2098 (SEQ ID NO:821). The NFAT response element may include nucleic acid NFAT consensus sequence of (5'-catttttccat-3') (SEQ ID NO:819) or (5'-tttttcca-3') (SEQ ID NO:820), which NFATc1 specifically binds to (Xu et al., Closely related T-memory stem cells correlate with in vivo expansion of CAR. CD19-T cells and are preserved by IL-7 and IL-15, Blood 2014 123:3750-3759), or repeats thereof. The NFAT response elements may also be separated by nucleic acid spacer sequences. Other NFAT responsive elements may exist and may further be discovered, and a skilled artisan in the art when directed to determine NFAT responsive element may do so by carrying out molecular biological assays to obtain it given the guidance of at least the responsive elements as set forth as SEQ ID NOS: 804-814 albeit as only mere examples. In one aspect of the invention, the cleavage enzyme that is downstream of the NFAT-response element and CMV minimal promoter is MMP9 (SEQ ID NO:652). In another aspect of the invention, the cleavage enzyme is a catalytic sub-unit of MMP9 (SEQ ID NO:653).

**[00564]** Because NFATs 1-4 are regulated by the calcineurin pathway, potential toxicities that may arise in a patient can be stopped by treatment with an immunosuppressive agent such as

FK506, Cyclosporin, Cyclosporin A, or Tacrolimus that block calcineurin activity and inhibit NFAT translocation to the nucleus. The T cell transduced or transfected with a cleavage enzyme on an inducible promoter may also be transduced or transfected with a CAR that recognizes a protein or molecule on the cancer cell. In a specific example, the cleavage enzyme is one that is able to cleave MUC1 full-length and the CAR bears an antibody fragment that directs it to MUC1\* on the surface of cancer cells.

**[00565]** To determine which cleavage enzymes cleave MUC1 on cancer cells, we tested a series of MMP and ADAM enzyme inhibitors. These experiments pointed to MMP9 as being an important cleavage enzyme in cancer cells. To confirm that MMP9 cleaves MUC1 on cancer cells, we transfected HCT-116 MUC1 negative colon cancer cells with a mimic of full-length MUC1 having 41 tandem repeat domains: HCT-MUC1-41TR. Through single cell cloning we were able to establish this cell line wherein MUC1 only minimally gets cleaved to MUC1\*. **Figs. 36A-36D** show Western blots and FACS analysis showing that HCT-MUC1-41TR is 95% positive for full-length MUC1 and only 5-10% positive for the cleaved form, MUC1\*. HCT-MUC1-41TR cells were incubated with MMP9 at varying concentrations and then assayed by immunofluorescence to measure binding of MNC2 monoclonal antibody to the resultant cells. As can be seen in **Figs. 37A-37C** binding of MNC2 increased as the concentration of MMP9 added to the cells increased. These experiments show that MMP9 cleaves MUC1 to a form that is recognized by MNC2. The human cancer tissue array studies we performed (Fig. 30A-30F, Fig. 31A-31F, Fig. 32A-32F, Fig. 33A-33F) show that MNC2 recognizes the form of cleaved MUC1 that is present on cancerous tissue but not on healthy cells or tissues (Fig. 34A-34I). Importantly, MNC2 does not recognize the form of cleaved MUC1 that is expressed on healthy hematopoietic stem cells of the bone marrow (Figs. 39-41).

**[00566]** In one aspect of the invention, an immune cell is transduced with both a CAR to target the immune cell to the tumor, and a cleavage enzyme. The CAR and the cleavage enzyme can be encoded on the same plasmid or on two different plasmids. In one aspect, the cleavage enzyme is on an inducible promoter. In another aspect, expression of the cleavage enzyme is induced by a protein that is expressed when the immune cell is activated. In one case, expression of the cleavage enzyme is induced by an NFAT protein. In another aspect, expression of the cleavage enzyme is induced by NFATc1. In another aspect, expression of the cleavage enzyme is induced when one of the NFAT proteins binds to an NFAT response element that is inserted

upstream of the gene for the cleavage enzyme or a catalytically active fragment thereof. In one aspect, the cleavage enzyme is MMP9 or a fragment of MMP9 that is catalytically active.

**[00567]** In one aspect of the invention, the cleavage enzyme is MMP9 (SEQ ID NO:643). Some cleavage enzymes are naturally expressed as pro-enzymes that need to be activated. This can be accomplished by biochemical means, by expressing a co-enzyme that activates a cleavage enzyme or by engineering the enzyme in an activated form. The invention anticipates overcoming this problem by co-expressing the cleavage enzyme with its activator. In one aspect of the invention, the cleavage enzyme is MMP9 and the co-activator is MMP3. In another aspect of the invention, the cleavage enzyme is expressed in a form that is already active, for example by expressing a fragment of the cleavage enzyme that still has catalytic function. In one case, the cleavage enzyme is an MMP9 fragment that is catalytically active. One example of an MMP9 catalytic fragment is given as SEQ ID NO:645.

**[00568]** MMP9, which must be activated by MMP3, is overexpressed in a large percentage of solid tumors. Further, it is known that MNC2 anti-MUC1\* monoclonal antibody recognizes MUC1 after it is cleaved by MMP9. The various breast, ovarian, pancreatic and lung cancer tissue arrays that were shown in **Figs. 30-33** were probed with MNC2-scFv, further indicating that MUC1 in these cancers is being cleaved by MMP9. To see if cleavage of tumors by MMP9 would increase T cell access to the tumor, we did a series of experiments using a cell line that expresses full-length MUC1, HCT-MUC1-41TR, a breast cancer cell line that is a high expresser of both full-length MUC1 and MUC1\* and a MUC1 negative cell line that we transfect with MUC1\*<sub>45</sub>. We transfected cells with MMP9 and MMP3, which activates MMP9. We took the supernatant of those cells, which contained activated MMP9, and added it to the various cells, which were then co-cultured with T cells transduced with an anti-MUC1\* CAR: huMNC2-CAR44. The result was greatly increased CAR T cell killing of the targeted MUC1/MUC1\* positive cancer cells, compared to the control cells that were not incubated with a MUC1 cleavage enzyme.

**[00569]** APMA is a biochemical that activates MMPs. We used APMA along with the conditioned media of cells that we transfected with either MMP9 or ADAM17 to see if any of these cleavage enzymes would cleave MUC1 on the HCT-MUC1-41TR cell line that only expresses full-length MUC1. As controls, we also tested the enzymes on HCT-MUC1\* cells. The MUC1 and MUC1\* expressing cells were stained with a red dye, CMTMR. Human T cells that

were transduced with an anti-MUC1\* CARs, CAR44 or CAR50 were co-cultured with the cancer cells. Untransduced T cells were used as a control (Fig. 45A-45P). As can be seen in **Fig. 45B**, **Fig. 45C**, and **Fig. 45D**, the anti-MUC1\* CAR T cells effectively recognized and clustered the HCT-MUC1\* cancer cells, which is a sign of T cell activation and killing. However, no CAR T cell induced clustering is visible in the wells containing HCT-MUC1-41TR, the full-length MUC1 expressing cells (**Fig. 45F**, **Fig. 45G**, and **Fig. 45H**). However, the cells that were incubated with activated MMP9 show dramatic increase in CAR T cell induced clustering (**Fig. 45J**, **Fig. 45K**, and **Fig. 45L**), indicating that MMP9 cleaved the full-length MUC1 to a form of MUC1\* that is recognized by MNC2 monoclonal antibody and more specifically by huMNC2-scFv. ADAM17 had no apparent effect. ADAM17 either did not cleave MUC1 or cleaved it at a position that is not recognized by MNC2, which is more likely (Fig. 45N-45P).

**[00570]** We performed the same experiment, this time using T47D breast cancer cells that were hard to kill using anti-MUC1\* CAR T cells presumably because they express high levels of full-length MUC1 as well as MUC1\* (Fig. 46A-46T). As can be seen in **Figs. 46B**, **46C**, and **46D**, anti-MUC1\* CAR44 and CAR50 have little effect on the T47D cancer cells. Only in **Fig. 46D**, which is CAR44 at the highest level of CAR expression in the T cells, do we see a small amount of CAR T cell induced clustering. However, the presence of activated MMP2 (**Fig. 46J**, **46K**, **46L**) or activated MMP9 (**Fig. 46R**, **46S**, **46T**) shows a dramatic increase in CAR T cell recognition, clustering and killing, showing that cleavage of full-length MUC1 increases T cell access to the cancer cells. To ensure that the addition of the APMA was not inducing cleavage or anti-MUC1\* CAR T recognition by some other mechanism, we made a catalytically active form of MMP9 and added it to T47D cells that were then co-cultured with MNC2-CAR44 T cells (Fig. 47A-47I). As can be seen in the figure, MNC2-CAR T cells recognize and cluster cells transfected with MUC1\* (Fig. 47B-47C), poorly cluster T47D breast cancer cells that express both full-length MUC1 and MUC1\* (Fig. 47E-47F), but robustly bind to and cluster the T47D cells after the addition of the catalytically active MMP9 (Fig. 47H-47I). This results supports the claim that MNC2 does not recognize full-length MUC1 but does recognize the growth factor receptor MUC1\*. Note that the full-length MUC1 expressed on this cell line may sterically hinder the binding of CAR T cells near the cell membrane.

**[00571]** In another example, T47D MUC1 positive tumor cells were incubated with a recombinant catalytic domain of MMP9 (Enzo Life Sciences, Inc., Farmingdale, NY) at either

100ng/mL or 500ng/mL. Western blot analysis showed that the MUC1/MUC1\* positive cancer cells underwent extensive cleavage of MUC1 to MUC1\*. In another example, T47D breast cancer cells were pre-incubated with a human recombinant MMP9 catalytic domain protein then co-cultured with anti-MUC1\* CAR44 T cells. The specific killing of the T47D cells by CAR44 T cells was monitored in real-time on an xCelligence instrument that measures impedance as a function of time. This analysis uses electrode arrays upon which cancer cells are plated. The adherent cancer cells insulate the electrode and cause an increase in impedance as they grow. Conversely, T cells are not adherent and remain in suspension so do not increase or decrease impedance. However, if the T cells or CAR T cells kill the cancer cells on the electrode plate, the cancer cells ball up and float as they die, which causes the impedance to decrease. The addition of MMP9 catalytic domain dramatically increased the killing of T47D cancer cells. **Fig. 48** shows an xCelligence graph of T47D breast cancer cells in co-culture with either untransduced T cells, as a control, or huMNC2-CAR44 T cells over a 45 hour period. After 18 hours of cancer cell growth, a catalytic sub-unit MMP9 was added to some of the cells. At 25 hours, T cells were added. As can be seen, huMNC2-CAR44 T cell killing is greatly improved when the T47D cells are pre-incubated with cleavage enzyme MMP9. In the xCelligence system, target cancer cells, which are adherent, are plated onto electrode array plates. Adherent cells insulate the electrode and increase the impedance. The number of adherent cancer cells is directly proportional to impedance. T cells are not adherent and do not contribute to impedance. Therefore, increasing impedance reflects growth of cancer cells and decreasing impedance reflects killing of cancer cells. Prostate cancer cell line DU145 expresses both MUC1 and MUC1\* but at a much lower level of expression than T47D cells. DU145 cells are efficiently killed by anti-MUC1\* CAR T cells in the presence or absence of a cleavage enzyme.

**[00572] Fig. 49** shows an xCelligence graph of DU145 prostate cancer cells in co-culture with either untransduced T cells, as a control, or huMNC2-CAR44 T cells over a 45 hour period. After 18 hours of cancer cell growth, a catalytic sub-unit MMP9 was added to some of the cells. At 25 hours, T cells were added. As can be seen, huMNC2-CAR44 T cell killing of low density MUC1/MUC1\* positive cancer cells is not affected by pre-incubation with cleavage enzyme MMP9. DU145 cancer cells express a significantly lower amount of MUC1 which includes the full-length form as well as MUC1\*. The lower density of full-length MUC1 does not sterically hinder T cell access to the membrane proximal MUC1\*. DU145 cells represent an early stage

cancer that expresses both full length and cleaved MUC1 but at lower levels so that T cell access is not sterically hindered. T47D cells represent mid-stage cancers that express high levels of both MUC1 and MUC1\*, wherein the density of MUC1 full-length sterically hinders access of T cells to the tumor. HCT-MUC1\* cells are a MUC1 negative cell line that has been stably transfected with MUC1\*<sub>45</sub>, and they represent late stage cancer cells. It is significant that MUC1 cleaved to MUC1\* by MMP9 is recognized by the anti-MUC1\* antibody MNC2, which is the targeting head of the CAR. Immune cell access to tumor antigens on the cancer cell surface can be sterically hindered by the presence of bulky extra cellular domain proteins or other obstructing elements also known as the tumor micro-environment. The aforementioned serve as an example that can be extended to improve the efficacy of CAR T therapies that target other tumor antigens. In one aspect of the invention, an immune cell is transfected or transduced with both a CAR comprising an antibody fragment that targets a tumor antigen and a cleavage enzyme. In another aspect of the invention, an immune cell is transfected or transduced with both a CAR comprising an antibody fragment that targets a tumor antigen and a cleavage enzyme that cleaves a tumor antigen to a form recognized by the antibody fragment of the CAR. In one aspect, an immune cell is transfected or transduced with both a CAR comprising an antibody fragment that targets a tumor antigen and a cleavage enzyme that cleaves a tumor antigen to a form recognized by the antibody fragment of the CAR, wherein the antibody fragment of the CAR recognizes MUC1\* extra cellular domain and the cleavage enzyme cleaves MUC1 to MUC1\*. In one aspect, an immune cell, which may be a T cell or an NK cell, is transfected or transduced with a CAR comprising an antibody fragment derived from MNC2, MNE6, MNC3 or MNC8 and a cleavage enzyme chosen from the group comprising MMP1, MMP2, MMP3, MMP7, MMP8, MMP9, MMP11, MMP12, MMP13, MMP14, MMP16, ADAM9, ADAM10, ADAM17, ADAM 19, ADAMTS16, ADAM28 or a catalytically active fragment thereof.

**[00573]** In one aspect of the invention, a person diagnosed with cancer or at risk of developing cancer is administered a sufficient amount of an immune cell transduced with both a CAR and a cleavage enzyme. In another aspect of the invention, a person diagnosed with cancer or at risk of developing cancer is administered a sufficient amount of an immune cell transduced with both a CAR and a cleavage enzyme, wherein the cleavage enzyme is on an inducible promoter that is activated by proteins that are expressed when the immune cell becomes activated. In another aspect of the invention, a person diagnosed with cancer or at risk of developing cancer is

administered a sufficient amount of an immune cell transduced with both a CAR and a cleavage enzyme, wherein the cleavage enzyme is on an inducible promoter that is activated by one or more NFAT. In one case the NFAT is NFATc1. In another aspect, the NFAT is NFATc3. In another aspect, the NFAT is NFATc2. In any of the instances above, the extra cellular domain of the CAR comprises a fragment of an anti-MUC1\* antibody. In one aspect, the anti-MUC1\* antibody is MNC2scFv or a humanized form of MNC2scFv. In another aspect, the anti-MUC1\* antibody is MNE6scFv or a humanized form of MNE6scFv. In any of the instances above, the immune cell can be a T cell, an NK cell, a mast cell, or a dendritic cell.

**[00574]** It is not intended that the present invention be limited to one or two specific methods of having expression of a cleavage enzyme induced by an activated T cell. We have demonstrated specific expression of a cleavage enzyme only upon T cell activation by constructing a plasmid with the cleavage enzyme gene downstream of an NFAT promoter sequence or downstream of one or more repeats of NFAT response elements. In another aspect of the invention, expression of the cleavage enzyme is induced by constructing a plasmid where the cleavage enzyme gene is inserted downstream of an IL-2 promoter sequence or downstream of an IL-2 response element, then inserting the plasmid into an immune cell. In another aspect of the invention, expression of the cleavage enzyme is induced by constructing a plasmid where the cleavage enzyme gene is inserted downstream of a Calcineurin promoter sequence or downstream of a Calcineurin response element, then inserting the plasmid into an immune cell and then administering to a patient for the treatment or prevention of cancers. There are also drug-inducible plasmids that can be used to induce expression of the cleavage enzyme or used to stop expression induced by an element of an activated T cell. These drug inducible systems may include tetracycline-inducible systems, Tet-on, Tet-off, tetracycline response elements, doxycycline, tamoxifen inducible systems, ecdysone inducible systems and the like.

**[00575]** It is not intended that the present invention be limited to one or two specific promoters used in the plasmids encoding the CARs or inducible cleavage enzymes. As is known by those skilled in the art, many promoters can be interchanged including SV40, PGK1, Ubc, CAG, TRE, UAS, Ac5, polyhedron, CaMKIIa, GAL1, GAL10, TEF1, GDS, ADH1, CaMV35S, Ubi, H1 and U6. Another solution to the problem of steric hindrance of CAR T cell access, caused by bulky cell surface proteins such as MUC1-FL, is to increase the length of the linker region of the CAR that is expressed by the T cell. In standard design CARs, the length of the

extracellular linker region between the transmembrane portion and the antibody fragment is about 45-50 amino acids in length. We made long-arm CARs where the length of the extracellular linker is extended from about 50 amino acids to 217 – 290 amino acids. Co-culture assays show that CARs with longer extracellular linkers have improved access to the tumor-associated antigen on the target cancer cells.

**[00576] BiTEs**

**[00577]** Divalent (or bivalent) single-chain variable fragments (di-scFvs, bi-scFvs) can be engineered by linking two scFvs. This can be done by producing a single peptide chain with two V<sub>H</sub> and two V<sub>L</sub> regions, yielding tandem scFvs. Another possibility is the creation of scFvs with linker peptides that are too short for the two variable regions to fold together (about five amino acids), forcing scFvs to dimerize. This type is known as diabodies. Diabodies have been shown to have dissociation constants up to 40-fold lower than corresponding scFvs, meaning that they have a much higher affinity to their target. Consequently, diabody drugs could be dosed much lower than other therapeutic antibodies and are capable of highly specific targeting of tumors *in vivo*. Still shorter linkers (one or two amino acids) lead to the formation of trimers, so-called triabodies or tribodies. Tetrabodies have also been produced. They exhibit an even higher affinity to their targets than diabodies.

**[00578]** All of these formats can be composed from variable fragments with specificity for two different antigens, in which case they are types of bispecific antibodies. The furthest developed of these are bispecific tandem di-scFvs, known as bi-specific T-cell engagers (BiTE antibody constructs). BiTEs are fusion proteins consisting of two scFvs of different antibodies, on a single peptide chain of about 55 kilodaltons. One of the scFvs may bind to T cells such as via the CD3 receptor, and the other to a tumor cell via a tumor specific molecule, such as aberrantly expressed MUC1\*.

**[00579]** Another aspect of the invention is a method for treating a patient diagnosed with, suspected of having, or at risk of developing a MUC1 positive or MUC1\* positive cancer, wherein the patient is administered an effective amount of a BiTE wherein one antibody variable fragment of the BiTE binds to a T cell surface antigen and the other antibody variable fragment of the BiTE binds to PSMGFR (SEQ ID NO:2), or more specifically to N-10 peptide. In one case, the antibody variable fragment of the BiTE that binds to MUC1\* comprises portions of



MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11.

**[00580]** In another aspect of the invention, MUC1\* peptides including PSMGFR (SEQ ID NO:2), or most or all of N-10 peptide are used in adoptive T cell approaches. In this case, a patient's T cells are exposed to the MUC1\* peptides and through various rounds of maturation, the T cells develop MUC1\* specific receptors. The adapted T cells are then expanded and administered to the donor patient who is diagnosed with, suspected of having, or is at risk of developing a MUC1\* positive cancer.

**[00581]** A series of CARs were also made that had MNC2 and humanized MNC2 as the extra cellular, targeting head of the CAR. The constructs for these CARs were inserted into a plasmid that was then inserted into a Lenti viral vector. Human T cells were then transduced with the lenti viral vector carrying the MNC2 CARs and huMNC2 CARs. MNC2-scFv-CARs that were mouse sequence or humanized were generated. In one aspect of the invention, the CAR comprised huMNC2-scFv-short hinge region-transmembrane domain derived from CD8-short intracellular piece-4-1BB-3zeta. In another aspect, the transmembrane domain was derived from CD4 transmembrane sequence. In another aspect, the intracellular co-stimulatory domain was CD28-3zeta. In yet another aspect, the intracellular co-stimulatory domain was CD28-4-1BB-3zeta.

**[00582]** There are a variety of methods for assessing whether or not T cells recognize a target cell and are in the process of mounting an immune response. T cells cluster when they recognize a target or foreign cell. This can be readily seen with the naked eye or at low magnification. The appearance of CAR T cell clustering when co-cultured with target cancer cells is one measure of: a) whether or not they recognize the cells as target cells; and b) whether or not they are getting activated to attack the targeted cells, which in this case are cancer cells. Figures 45-47 show photographs of MUC1\* positive T47D breast cancer cells that were either stably transfected with mCherry or dyed with CMTMR, so are red, which were co-cultured with either human T cells without a CAR or human T cells transduced with huMNC2-scFv-CAR44, or with huMNC2-scFv-CAR50. The CAR T cells are clear. As can be seen, there is no T cell induced clustering of the cancer cells when the T cell does not carry a CAR. However, when T cells carrying a MUC1\* targeting CAR, there is dramatic clustering of the MUC1\* positive cancer cells.

**[00583]** After T cells recognize and cluster target cells, they overexpress perforin and granzyme B. Together these two molecules activate a cell death pathway in the targeted cell. It is thought that the perforin makes a hole in the target cell into which the T cell injects granzyme B which then activates apoptotic proteases, causing the target cell to lyse. **Figs. 55 and Fig. 56** show huMNC2-scFV-CAR44 T cells binding to target MUC1\* positive prostate cancer and pancreatic cancer cells and injecting granzyme B.

**[00584]** Another measure of whether or not a T cell has recognized a target cell and is activated to kill that cell, is the upregulation and secretion of cytokines, interferon gamma (IFN-g) and interleukin-2 (IL-2), by the T cell. Activation of CAR T cells, as evidenced by IFN-g and IL-2 secretion, can be readily measured *in vitro*. CAR T cells are co-cultured with target cells and after an incubation period, the conditioned media is assayed by ELISA to detect secreted IFN-g and IL-2. In order to determine the cancer-specificity of CAR T cells wherein the targeting head of the CAR was either huMNC2 or huMNE6, these experiments were performed with huMNC2-CAR44 T cells and huMNE6-CAR44 T cells in co-culture with MUC1\* positive cancer cells and normal cells. Table 1 details the MUC1 positive normal or primary cells that were tested.

**[00585] Table 1: Normal Cell Lines and Primary Cells**

**[00586]**

Cell Line	ATCC Designation	Tissue	Origin
Hep.G2		Liver	
THLE-3	CRL-11233	Liver	The THLE-2 (ATCC CRL-10149 and the THLE-3 (ATCC CRL-11233) cell lines were derived from primary normal liver cells by infection with SV40 large T antigen. THLE-2 and THLE-3 cells express phenotypic characteristics of normal adult liver epithelial cells. They are nontumorigenic when injected into athymic nude mice, have near-diploid karyotypes, and do not express alpha-fetoprotein.

Cell Line	ATCC Designation	Tissue	Origin
Lonza Primary Hepatocytes	HUM181141	Liver	Male, Caucasian 2.0 months old Induction Fold CYP1A2 (a) 14.0 Induction Fold CYP2B6 (b) 13.0 Induction Fold CYP3A4 (c) 44.0 Basal Activity CYP1A2 2.6 Basal Activity CYP2B6 0.7 Basal Activity CYP3A4 14.0 Additional Information: Inducer / Marker Metabolite (a) 0.05 mM Omeprazole / Acetaminophen (b) 1 mM Phenobarbital / Hydroxybupropion (c) 0.01 mM Rifampicin / 6-Beta-Hydroxytestosterone Basal activity is expressed as: pmol/million cells/minute
T/G HA-VSMC	CRL-1999	Aortic Smooth Muscle	11 months Female, Caucasian
CCD-18Lu	CCL-205	Lung	This fibroblast-like cell line was derived from the lung tissue of a <b>2 month, 17-day-old Black female</b> . The donor had cerebral anoxia, cardiac anomaly, sepsis, endocardial cushion defect and fetal alcoholic syndrome. Female, Black 2.5 months
HBEC-5i	CRL-3245	Brain endothelium	Derived from small fragments of human cerebral cortex obtained from patients who had died of various causes.
Hs 738.St/Int	CRL-7869	Stomach/Intestine	<b>18 weeks gestation fetus</b> Male, Caucasian Part of the NBL Cell Line Collection. This cell line is neither produced nor fully characterized by ATCC. We do not guarantee that it will maintain a specific morphology, purity, or any other property upon passage.
MCF-12A	CRL-10782	Breast	The MCF-12A cell line is a non-tumorigenic epithelial cell line established from tissue taken at reduction mammoplasty from a nulliparous patient with fibrocystic breast disease that contained focal areas of intraductal hyperplasia. The line was produced by long term culture in serum free medium with low Ca++ concentration. MCF-12A was

Cell Line	ATCC Designation	Tissue	Origin
			derived from adherent cells in the population.
Hs 1.Tes	CRL-7002	Testis	Male, Caucasian <b>second trimester</b> Part of the NBL Cell Line Collection. This cell line is neither produced nor fully characterized by ATCC. We do not guarantee that it will maintain a specific morphology, purity, or any other property upon passage.
HRCE	Lonza: catalogue # CC-2554 Lot # 0000542104	Kidney	Human Renal Cortical Cells (HRCE) are from proximal and distal tubules. Donor info: 49 year old female, passage 2, 95% viability, doubling time (hours) 24 hrs

[00587]

[00588] Figure 50 is a graph of PCR measurement of the various cell lines tested, wherein mRNA levels of MUC1 are measured. The cancer cell lines that were tested in these assays were HCT-MUC1\* and T47D breast cancer cells. These cells were co-cultured with huMNC2-CAR44 human T cells. Co-culture of huMNC2-CAR44 T cells with the cancer cells induced the CAR T cells to secrete large amounts of IFN-g and IL-2 into the surrounding media, yet co-culture with the MUC1 positive normal cells induced no secretion of the cytokines (Fig. 51 and Fig. 52). In addition to testing for IFN-g and IL-2 secretion by the CAR T cells, the normal cells were assayed for signs of cell death, which could have been induced by the CAR T cells if the antibody targeting head were not extremely cancer-specific. After co-culture with huMNC2-CAR44 T cells, the cells were incubated with a cell death marker, then assayed by FACS. huMNC2-CAR44 T cells induced no cell death in the normal cells (Fig. 53A-53J).

[00589] In addition to FACS analysis, many researchers now use an xCELLigence instrument to measure CAR T killing of cancer cells. FACS is not the best method for tracking T cell induced cell killing because the T cells lyse the target cell. By FACS it is difficult to measure

dead cells because they are excluded as cell debris, so one must infer an amount of cell killing and by various methods determine if the missing cells are T cells or cancer cells.

**[00590]** The xCELLigence instrument uses electrode arrays upon which cancer cells are plated. The adherent cancer cells insulate the electrode and so cause an increase in impedance as they grow. Conversely, T cells are not adherent and remain in suspension so do not contribute to insulation of the electrode which would increase impedance. However, if the T cells or CAR T cells kill the cancer cells on the electrode plate, the cancer cells ball up and float off as they die, which causes the impedance to decrease. The xCELLigence instrument measures impedance as a function of time, which is correlated to cancer cell killing. In addition, the electrode plates also have a viewing window. When CAR T cells effectively kill the adsorbed target cancer cells, there is a decrease in impedance but also one can see that there are no cancer cells left on the plate surface.

**[00591]** **Figs. 55A-55H** show the cytotoxic effect of huMNC2-CAR44 T cells on MUC1\* positive DU145 prostate cancer cells as measured by a variety of assays. **Fig. 55A** is a fluorescent photograph of untransduced T cells co-cultured with the prostate cancer cells, wherein granzyme B is stained with a red fluorophore. **Fig. 55C** is a fluorescent photograph of huMNC2-CAR44 T cells co-cultured with the prostate cancer cells, wherein granzyme B is stained with a red fluorophore. **Fig. 55D** is the DAPI and granzyme B merge. **Fig. 55E** is a FACS scan for fluorescently labeled granzyme B for untransduced T cells incubated with the cancer cells. **Fig. 55F** is a FACS scan showing a positive increase in fluorescently labeled granzyme B for huMNC2-CAR44 T cells incubated with the cancer cells. **Fig. 55G** is a graph of the mean fluorescent intensity. **Fig. 55H** is an xCELLigence scan tracking the real-time killing of DU145 cancer cells by huMNC2-CAR44 T cells (blue trace) but not by untransduced T cells (green). **Figs. 56A-56H** show the cytotoxic effect of huMNC2-CAR44 T cells on MUC1\* positive CAPAN-2 pancreatic cancer cells as measured by a variety of assays. **Fig. 56A** is a fluorescent photograph of untransduced T cells co-cultured with the pancreatic cancer cells, wherein granzyme B is stained with a red fluorophore. **Fig. 56B** is the DAPI and granzyme B merge. **Fig. 56C** is a fluorescent photograph of huMNC2-CAR44 T cells co-cultured with the pancreatic cancer cells, wherein granzyme B is stained with a red fluorophore. **Fig. 56D** is the DAPI and granzyme B merge. **Fig. 56E** is a FACS scan for fluorescently labeled granzyme B for untransduced T cells incubated with the cancer cells. **Fig. 56F** is a FACS scan showing a positive

increase in fluorescently labeled granzyme B for huMNC2-CAR44 T cells incubated with the cancer cells. **Fig. 56G** is a graph of the mean fluorescent intensity. **Fig. 56H** is an xCELLigence scan tracking the real-time killing of CAPAN-2 cancer cells by huMNC2-CAR44 T cells (blue trace) but not by untransduced T cells (green). **Figs. 57A-57C** show xCELLigence scans tracking the real-time killing of MUC1\* positive cancer cells, but not MUC1\* negative cells, by huMNC2-CAR44 T cells. **Fig. 57A** shows that huMNC2-CAR44 T cells effectively kill HCT colon cancer cells that have been stably transfected with MUC1\*. **Fig. 57B** shows that huMNC2-CAR44 T cells have almost no effect on HCT-MUC1-41TR, which is a MUC1 negative cancer cell that has been stably transfected with a MUC1 full-length. In this cell line only about 10% of the cell have MUC1 cleaved to MUC1\*. **Fig. 57C** shows that huMNC2-CAR44 T cells have no effect on HCT-116 cells, which is a MUC1 negative colon cancer cell line.

**[00592]** These data demonstrate that T cells transduced with a CAR wherein the antibody fragment targeting head is MNC2, effectively kill MUC1\* positive cancer cells. These data specifically show that huMNC2-scFV-CAR44 transduced into human T cells effectively kill MUC1\* positive cancer cells. Because we and others have now demonstrated that the most important aspect of CAR T function is the targeting antibody fragment, it follows that an immune cell or a T cell transduced with any CAR having the antibody fragment MNC2-scFV or huMNC2-scFV would have similar efficacy against MUC1 or MUC1\* positive tumors. For example, the hinge region that connects the scFv to the transmembrane portion could be any flexible linker. The intracellular co-stimulatory domains could be CD28-3zeta, CD28-4-1BB-3zeta or any combination of immune cell co-stimulatory domains.

**[00593]** Figure 61 shows an experiment in which huMNC2-scFv-CAR44 transduced human T cell that were bead stimulated (Protocol 1) or cancer cell stimulated (Protocol 2) were tested for their ability to inhibit tumor growth in animals. Human cancer cells that had been stably transfected with Luciferase were injected into female NOD/SCID/GAMMA (NSG) mice between 11 and 15 weeks of age. 500,000 BT-20 breast cancer cells were injected subcutaneously into a rear flank. Tumor engraftment was verified by injecting the animals with Luciferin and then imaging the fluorescent cancer cells using an IVIS instrument. IVIS images taken Day 5 post implantation showed the presence of tumor cells. On Day 6 after IVIS measurement, animals were given a one-time injection of 10 million of either human T cells transduced with huMNC2-scFv-CAR44 or untransduced T cells. 5 million T cells were injected

intra-tumor and 5 million were injected into the tail vein. 10 minutes prior to IVIS photographs, mice were IP injected with Luciferin, which fluoresces after cleavage by Luciferase, thus making tumor cells fluoresce. Figs. 61A, 61D, 61G show photographs of mice that were treated with huMNC2-scFv-CAR44 T cells that had been pre-stimulated by co-culturing for 24 hours with 4 $\mu$ m beads to which was attached a synthetic MUC1\*, PSMGFR peptide 24 hours prior to administration, "Protocol 1". Figs. 61B, 61E, 61H show photographs of mice that were treated with huMNC2-scFv-CAR44 T cells that had been pre-stimulated by twice co-culturing for 24 hours with MUC1\* positive cancer cells 24 hours prior to administration, "Protocol 2". As can be seen in **Figure 61**, huMNC2-CAR44 T cells that were peptide-bead stimulated inhibited tumor growth better than cells pre-stimulated by incubation with live cancer cells, which likely contaminated the target cells and increased the tumor volume.

**[00594]** huMNC2-scFv-CAR44 transduced human T cell that were bead stimulated (Protocol 1) or cancer cell stimulated (Protocol 2) were also tested for their ability to inhibit tumor growth in animals. Human cancer cells that had been stably transfected with Luciferase were injected into female NOD/SCID/GAMMA (NSG) mice between 11 and 15 weeks of age. In another experiment, 500,000 BT-20 MUC1\* positive triple negative breast cancer cells were injected sub-cutaneously into a rear flank. Tumor engraftment was verified by injecting the animals with Luciferin and then imaging the fluorescent cancer cells using an IVIS instrument. IVIS images taken Day 6 post implantation showed the presence of tumor cells. On Day 6, after IVIS imaging, 10M huMNC2-scFv-CAR44 T cells were administered to the animals. 5M of the CAR T cells were administered by intratumor injection and the other 5M were administered by tail vein injection. Control group was injected by same administration routes with the same number of untransduced T cells. IVIS measurements of tumor burden were taken on Days 6, 8, and 12. As can be seen in **Figs. 61A-61J**, both groups of mice treated with huMNC2-CAR44 T cells showed a decrease in tumor burden compared to the control group.

**[00595]** huMNC2-scFv-CAR44 transduced human T cell that were bead stimulated (Protocol 1) were also tested for their ability to inhibit ovarian cancer growth in animals. Human SKOV-3 MUC1\* positive ovarian cancer cells that had been stably transfected with Luciferase were injected into female NOD/SCID/GAMMA (NSG) mice between 11 and 15 weeks of age. In one experiment, 500,000 SKOV-3 cancer cells were injected into the intraperitoneal cavity to mimic metastatic ovarian cancer in humans. Tumor engraftment was verified by injecting the animals

with Luciferin and then imaging the fluorescent cancer cells using an IVIS instrument. IVIS images taken Day 3 post implantation showed the presence of tumor cells. On Day 4 and Day 11, post tumor implantation, 10M huMNC2-scFv-CAR44 T cells were IP administered to the animals. On Day 4, CAR T cells were IP injected. On Day 11 half the CAR T cells were injected into the intraperitoneal space and the other half was injected into the tail vein. Control groups were injected by same administration routes with either the same number of untransduced T cells or same volume of PBS. Subsequent IVIS measurements of tumor burden were taken on Day 7, Day 10 and Day 15. As can be seen in **Figs. 62A-62L**, control mice have tumors that are growing at a much faster rate than the huMNC2-CAR44 T cell treated mice. **Fig. 62M** shows the IVIS color bar correlating photons/second to color.

**[00596]** One aspect of the invention is a method for treating a patient diagnosed with, suspected of having, or at risk of developing a MUC1 positive or MUC1\* positive cancer, wherein the patient is administered an effective amount of immune cells that have been transduced with a MUC1\* targeting CAR, wherein the CAR is chosen from among the group consisting of MN-E6-CD8-CD28-3z (SEQ ID NOS:297-298); MN-E6-CD4-CD28-3z (SEQ ID NOS:748-749); MN-E6-CD8-41BB-3z (SEQ ID NOS:300-301); MN-E6-CD4-41BB-3z (SEQ ID NOS:750-751); MN-E6-CD8-CD28-41BB-3z (SEQ ID NOS:303-304); MN-E6-CD4-CD28-41BB-3z (SEQ ID NOS:754-755); MN-E6scFv-Fc-8-41BB-CD3z (SEQ ID NOS:310-311); MN-E6scFv-IgD-Fc-8-41BB-CD3z (SEQ ID NOS:770-771); MN-E6scFv-FcH-8-41BB-CD3z (SEQ ID NOS:315-316); MN-E6scFv-IgD-FcH-8-41BB-CD3z (SEQ ID NOS:772-773); MN-E6scFv-Fc-4-41BB-CD3z (SEQ ID NOS:318-319); MN-E6scFv-FcH-4-41BB-CD3z (SEQ ID NOS:321-322); MN-E6scFv-IgD-8-41BB-CD3z (SEQ ID NOS:323-324); MN-E6scFv-IgD-4-41BB-CD3z (SEQ ID NOS:327-328); MN-E6scFv-X4-8-41BB-CD3z (SEQ ID NOS:330-331); MN-E6scFv-X4-4-41BB-CD3z (SEQ ID NOS:333-334); MN-E6scFv-8-4-41BB-CD3z (SEQ ID NOS:336-337), or any of the aforementioned CARs wherein the MN-E6 is replaced by fragment derived from MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11. Another aspect of the invention is a method for treating a patient diagnosed with, suspected of having, or at risk of developing a cancer, wherein the patient is administered an effective amount of immune cells that have been transduced with one of the aforementioned CARs wherein the MN-E6 is replaced by a peptide comprising antibody variable domain



fragments that are specific for a cancer antigen. In any of the above methods, the immune cell may be a T cell and may further be isolated from the patient to be treated.

**[00597] Other MUC1 cleavage sites**

**[00598]** It is known that MUC1 is cleaved to the growth factor receptor form, MUC1\*, on some healthy cells in addition to cancer cells. For example, MUC1 is cleaved to MUC1\* on healthy stem and progenitor cells. A large percentage of bone marrow cells are MUC1\* positive. Portions of the intestine are MUC1\* positive.

**[00599]** The inventors have discovered that MUC1 can be cleaved at different positions that are relatively close to each other but the location of cleavage changes the fold of the remaining portion of the extracellular domain. As a result, monoclonal antibodies can be identified that bind to MUC1\* cleaved at a first position but do not bind to MUC1\* that has been cleaved at a second position. This discovery is disclosed in WO2014/028668, filed August 14, 2013, the contents of which are incorporated by reference herein its entirety. We identified a set of anti-MUC1\* monoclonal antibodies that bind to MUC1\* as it appears on cancer cells but do not bind to MUC1\* as it appears on stem and progenitor cells. Conversely, we identified a second set of monoclonal antibodies that bind to stem and progenitor cells but do not bind to cancer cells. One method used to identify stem specific antibodies is as follows: supernatants from monoclonal hybridomas were separately adsorbed onto 2 multi-well plates. Stem cells, which are non-adherent cells, were put into one plate and cancer cells which are adherent were put into an identical plate. After an incubation period, the plates were rinsed and inverted. If the non-adherent stem cells stuck to the plate, then the monoclonal antibody in that particular well recognizes stem cells and will not recognize cancer cells. Antibodies that did not capture stem cells or antibodies that captured cancer cells were identified as cancer specific antibodies. FACS analysis has confirmed this method works.

**[00600]** Antibodies MN-E6 and MN-C2 are examples of cancer-specific antibodies. Antibodies MN-C3 and MN-C8 are examples of stem-specific antibodies. Although both sets of antibodies are able to bind to a peptide having the PSMGFR sequence, FACS analysis shows that the anti-MUC1\* polyclonal antibody and MN-C3 bind to MUC1\* positive bone marrow cells but MN-E6 does not. The MUC1\* polyclonal antibody was generated by immunizing a rabbit with the PSMGFR peptide. Similarly, MN-C3 binds to stem cells of the intestinal crypts but MN-E6 does not. Conversely, MN-E6 antibody binds to cancerous tissue while the stem-

specific MN-C3 does not. Competition ELISA experiments indicate that the C-terminal 10 amino acids of the PSMGFR peptide are required for MN-E6 and MN-C2 binding, but not for MN-C3 and MN-C8. Therefore, another method for identifying antibodies that are cancer specific is to immunize with a peptide having the sequence of the PSMGFR peptide minus the 10 N-terminal amino acids or use that peptide to screen for antibodies or antibody fragments that will be cancer specific. Antibodies that bind to a peptide with a sequence of PSMGFR peptide minus the N-terminal 10 amino acids, referred to herein as N-10 peptide, but do not bind to a peptide with a sequence of PSMGFR peptide minus the C-terminal 10 amino acids, C-10 peptide, are cancer specific antibodies for use in the treatment or prevention of cancers.

**[00601]** The extracellular domain of MUC1 is also cleaved on stem cells and some progenitor cells, where activation of cleaved MUC1 by ligands NME1 in dimer form or NME7 promotes growth and pluripotency and inhibits differentiation. The transmembrane portion of MUC1 that remains after cleavage is called MUC1\* and the extracellular domain is comprised essentially of the Primary Sequence of MUC1 Growth Factor Receptor (PSMGFR) sequence. However, the exact site of cleavage can vary depending on cell type, tissue type, or which cleavage enzyme a particular person expresses or overexpresses. In addition to the cleavage site that we previously identified which leaves the transmembrane portion of MUC1\* comprising most or all of the PSMGFR (SEQ ID NO:2), other cleavage sites could possibly result in an extended MUC1\* comprised of most or all of SNIKFRPGSVVVQLTLAFREGTINVHDTVETQFNQYKTEAASRY (SEQ ID NO:620); or

**[00602]** SVVVQLTLAFREGTINVHDTVETQFNQYKTEAASRY (SEQ ID NO:621).

**[00603]** To test this hypothesis, and to determine if antibodies to an N-terminally extended PSMGFR, would generated more cancer-specific antibodies than antibodies that bind to the PSMGFR, we generated monoclonal antibodies by immunization with peptides:

**[00604]** (PSMGFR) GTINVHDTVETQFNQYKTEAASRYNLTISDVSVDVPFPFSAQSGA (SEQ ID NO:2),

**[00605]** (N+20/C-27) SNIKFRPGSVVVQLTLAFREGTINVHDTVETQFNQYKTE (SEQ ID NO:822), or

**[00606]** (N+9/C-9) VQLTLAFREGTINVHDTVETQFNQYKTEAASRYNLTISDVSVDVP (SEQ ID NO:824)

**[00607]** Monoclonal antibodies generated from immunization with the same peptide can also show differences in reactivity to the same cancerous tissue specimen. These results indicate that the monoclonal antibodies recognize different conformations of the truncated MUC1 extra cellular domain produced by immunizing with different length peptides, mimicking different cleavage sites, or from cleavage at different sites in the host animal. Antibodies that recognize different cleavage site conformations may be cancer sub-type specific or patient specific, depending on which cleavage enzyme their tumor expresses. In one aspect of the invention, a patient diagnosed with a certain type of cancer is treated with an antibody of the invention that recognizes a cleaved MUC1 wherein the antibody is specific for cleavage by a specific enzyme that is known to be typically expressed by that sub-type of cancer. In another aspect, a patient tumor is analyzed to determine which enzyme his or her tumor expresses and an antibody that recognizes a MUC1 cleaved by that enzyme is then administered to the patient for the treatment of their cancer. The antibody may be in the form of a CAR, a BiTE, an ADC, or a bispecific antibody.

**[00608]** We previously reported that it is the MUC1 transmembrane cleavage product, called MUC1\* (muk1 star), that mediates tumor growth and not full-length MUC1 (Mahanta et al 2008). MUC1\* is a growth factor receptor that is activated by ligand induced dimerization of its short extra cellular domain (Fig. 1A). Dimerization of the MUC1\* extra cellular domain activates the MAP kinase signaling cascade and stimulates growth and survival of cancer cells (Fessler et al 2009). Bivalent antibodies that dimerize the MUC1\* extra cellular domain stimulate cancer cell growth while the monovalent Fab of the same antibody, which cannot dimerize, inhibits cancer cell growth. We demonstrated this in vitro (Fig. 1B) and in vivo (Fig. 7A-7B ).

**[00609]** We then identified the natural ligands that dimerize and activate MUC1\* growth factor receptor function. Dimers of NME1 bind to and dimerize the MUC1\* extra cellular domain and stimulate growth (Fig. 1C and Smagghe et al 2013). NME1 can turn its growth factor properties off. NME1 is secreted by MUC1\* positive cells. Dimeric NME1 binds to MUC1\* to stimulate growth. However, as the cell population grows, more and more NME1 is secreted from the cells. At high concentrations, the NME1 dimers multimerize and form hexamers, which do not bind to MUC1\*, but likely bind to some unknown receptor, as the addition of NME1 hexamers turns off growth. NME1 is an adult form. The embryonic form is

NME7<sub>AB</sub> (Carter et al 2016). Each NME7<sub>AB</sub> monomer has two binding sites for MUC1\* so as a monomer it dimerizes MUC1\* (Fig. 1D), stimulates growth and cannot turn itself off. In the developing embryo, BRD4 turns off NME7 and its co-factor JMJD6 turns on the self-regulating form, NME1. However, in cancers, NME7, which should be silenced in adult life, is aberrantly expressed again, where it renders the MUC1\* growth factor receptor constitutively active.

**[00610]** *In vitro*, NME1 (SEQ ID NO:4) and NME7<sub>AB</sub> (SEQ ID NO:827) bind to the PSMGFR portion of the MUC1\* extra cellular domain. Both growth factors can bind to the PSMGFR peptide (SEQ ID NO:2) even if the 10 N-terminal amino acids are deleted, referred to herein as N-10 (SEQ ID NO:3). However, neither NME1 nor NME7<sub>AB</sub> can bind to the PSMGFR peptide if the 10 membrane proximal amino acids are deleted (Fig. 2A-2D), referred to herein as C-10 (SEQ ID NO:825). In summary, the epitope to which NME1 and NME7<sub>AB</sub> bind includes all or part of the 10 membrane proximal amino acids: PFPFSAQSGA (SEQ ID NO:1743). We tested various antibodies that were generated in animals by immunizing with the PSMGFR peptide for their ability to recognize cancer cells but not healthy cells. Among the most cancer selective were the MNC2 and MNE6 monoclonal anti-MUC1\* antibodies. Two other monoclonal antibodies that were generated from immunizing animals with the PSMGFR peptide are MNC3 and MNC8. Although MNC2, MNE6, MNC3 and MNC8 all bind to the PSMGFR peptide, like NME1 and NME7<sub>AB</sub>, MNC2 and MNE6 bind strongly to the N-10 peptide but not to the C-10 peptide (Fig. 2B-2C). In fact, MNC2 and MNE6 competitively inhibit the binding of NME1 and NME7<sub>AB</sub> to PSMGFR (Fig. 3A-3C). Conversely, MNC3 and MNC8 bind to the C-10 peptide, bind less well to the N-10 peptide and do not compete with NME1 nor NME7<sub>AB</sub> for binding to MUC1\* peptides, including PSMGFR (Fig. 2E-2F). MNC3 and MNC8 are far less cancer specific than MNC2 and MNE6. MNC3 and MNC8 recognize stem and progenitor cells, such as hematopoietic stem cells, whereas MNC2 and MNE6 do not (Fig. 39-41). Because hematopoietic stem cells are the progenitor cells for the blood cells, it would be problematic to have a cancer therapeutic that would also target such an important normal cell type.

**[00611]** Because MUC1\* is generated by enzymatic cleavage of MUC1, we researched which cleavage enzymes cleave MUC1 to a MUC1\* and whether or not we could identify antibodies that would recognize a MUC1\* generated by a first cleavage enzyme but not MUC1\* generated by a second cleavage enzyme. We found that MNC2 and MNE6 recognized a MUC1\* generated by cleavage of MUC1 by MMP9 but not by cleavage by other enzymes such as MMP2 (Fig. 37

and Fig. 75). We note that MMP9 is overexpressed in cancers and is a predictor of poor prognosis (vant Veer et al 2002; Dufour et al 2011) and has been implicated in metastasis (Owyong et al, 2019), whereas MMP2 is expressed in bone marrow. One antibody binding to a MUC1\* generated by cleavage by a first enzyme but not by cleavage by a second enzyme implies that the antibody recognizes a conformational epitope rather than a linear epitope.

**[00612]** We reasoned that the most cancer specific antibodies would be those antibodies that are characterized by some combination of most or all of the following:

**[00613]** Antibody binds to PSMGFR peptide;

**[00614]** Antibody does not bind to full-length MUC1;

**[00615]** Antibody binds to N-10;

**[00616]** Antibody does not bind to C-10;

**[00617]** Antibody competitively inhibits binding of NME1 or NME7<sub>AB</sub> to MUC1\* extra cellular domain or a PSMGFR peptide;

**[00618]** Antibody recognizes a MUC1\* generated by cleavage by MMP9;

**[00619]** Antibody recognizes a conformational epitope not a linear epitope.

**[00620]** **MNC2 and MNE6 are cancer specific.**

**[00621]** Our experiments show that both MNC2 and MNE6: a) Bind to tumor cells; b) monovalent forms block tumor growth in vitro and in vivo; c) have minimal to no binding of normal tissue while having robust binding to a wide panel of tumor tissues; d) when incorporated into CAR T cells, MNC2 and MNE6 directed CAR T cells do not recognize full-length MUC1 and do not kill cells that only express full-length MUC1; e) MNC2 and MNE6 directed CAR T cells cluster then kill tumor cells expressing MUC1\*; and f) MNC2 and MNE6 recognize a MUC1 cleavage product when it is cleaved by MMP9.

**[00622]** MNC2 directed CAR T cells do not recognize normal, healthy cells that are MUC1\* positive. A panel of normal cell lines, as well as primary cells, were co-cultured with huMNC2-CAR44 T cells. The normal cell populations were analyzed to determine whether or not the MNC2 directed CAR T cells killed them. The CAR T cells were analyzed to see if co-culture with the MUC1 positive normal cells activated the killing function of the CAR T cells, as measured by secretion of IL-2 or interferon gamma. As **Figures 50-52** show, the MNC2 directed CAR T cells did not kill the normal cells, nor was there cytokine secretion, indicative of T cell activation. In addition, over 2,000 human tissue specimens were analyzed. The results showed

that neither MNC2 nor MNE6 showed any significant binding to normal tissues but showed robust staining of a wide panel of cancerous tissues. For example, MNC2 stained 93% breast cancer specimens, 83% ovarian, 78% pancreatic and 71% lung cancer specimens. In addition, patient-matched primary tumors (Fig. 54) and subsequent metastases showed that the amount of MNC2-reactive MUC1\* increased with tumor progression and metastases. In summary, MNC2 is a highly cancer specific antibody.

**[00623] Characterization of MNC2 and MNE6**

**[00624]** Our gold standard, cancer-specific antibodies MNC2 and MNE6: 1) bind to N-10 peptide but not to the C-10 peptide; 2) compete with NME7<sub>AB</sub> and dimeric NME1 for the same binding site near the C-terminus of the PSMGFR peptide, which is the membrane proximal portion of MUC1\* on cells; 3) do not work in a Western blot assay indicating that they recognize a conformational rather than linear epitope; 4) recognize a MUC1\* generated when MUC1 is cleaved by MMP9; 5) do not bind to full-length MUC1 but only to the cleaved form, MUC1\*, in model cell lines as well as cancer cell lines; 6) show little to no binding to normal tissues but robustly stain a wide variety of tumor tissues; and 7) share some consensus sequences in their Complementarity Determining Regions, CDRs.

**[00625]** In an effort to identify other antibodies that are highly cancer-specific, like MNC2 and MNE6, we subjected new antibodies to a set of seven (7) characterization experiments: 1) epitope binding assays; 2) functional assays such as the ability to displace activating growth factor NME7<sub>AB</sub> or dimeric NME1 from binding to MUC1\* peptides PSMGFR or N-10; 3) Western blots to determine whether or not the antibodies recognized a linear epitope versus a conformational epitope, in which case the antibodies would not work in a Western; 4) binding assays to see if the antibodies recognized a cleaved MUC1 that was dependent on cleavage by MMP9; 5) FACS analysis to measure the ability of the antibodies to recognize MUC1\* positive cells but not full-length MUC1; and FACS analysis to measure the ability of the antibodies to recognize MUC1/MUC1\* positive cancer cells; 6) immunohistochemistry, IHC, assays of normal tissues versus cancerous tissues to determine true cancer specificity; and 7) aligning antibody sequences to determine if subsets of antibodies shared consensus sequences that could predict their cancer specificity or lack thereof.

[00626] Monoclonal antibodies were produced by immunizing animals with peptides derived from a MUC1 that is devoid of tandem repeats. These antibodies included PSMGFR and peptides that were extended at the N-terminus of PSMGFR. Immunizing peptides were:

[00627] PSMGFR (SEQ ID NO:2))

[00628] N+9/C-9 (9 amino acids added onto the N-terminus and 9 amino acids deleted from the C-terminus) (SEQ ID NO:824)

[00629] N+20/C-27 (20 amino acids added onto the N-terminus and 27 amino acids deleted from the C-terminus) (SEQ ID NO:823)

[00630] These monoclonal antibodies were then tested to determine which satisfied the seven (7) characterization criteria cited above, which we reasoned would identify the most cancer specific antibodies.

[00631] **Epitope binding assays**

[00632] ELISA assays were performed to determine if, in addition to recognizing their immunizing peptide, they recognized PSMGFR, N-10 or C-10. In addition, they were tested for their ability to bind to N+20/C-27, N+9/C-9. We first did the ELISA assay on our set of reference antibodies, MNC2, MNE6, which we know are cancer-specific plus MNC3, which we know recognizes stem cells and progenitor cells (Fig. 63A-63B). None of the reference antibodies bound to the N+20/C-27 peptide. MNC2 and MNE6 cannot bind to PSMGFR peptides with 27, 10 or 9 C-terminal deletions, however, MNC3 binds to C-10 and to N+9/C-9 peptides.

[00633] This same ELISA assay was performed on the antibodies of the invention (Fig. 64-66 and Fig. 201). The binding patterns of the antibodies that were generated by immunizing with the PSMGFR peptide are shown in Fig. 64A-64B. Note that only 20A10 exactly matches the binding profile of MNC2 and MNE6. 25E6, 28F9 and 18G12 are all able to bind to the N-10 peptide. 18B4 is the only antibody raised against the PSMGFR peptide that requires the 10 most N-terminal amino acids of the peptide. The color of the bars for each antibody in the ELISA graph are color coded to match the deductive cognate sequence, or a portion thereof, of that antibody. In addition, another set of antibodies was assayed by ELISA (Fig. 201). Of this set, B12, B2, B7, B9, 8C7F3, and H11 bound to the PSMGFR peptide, bound to the N-10 peptide, but not to the C-10 peptide (Fig. 201). The binding patterns of the antibodies that were generated by immunizing with the N+20/C-27 peptide are shown in **Fig. 65A-65B**. Although these antibodies were raised

against the N+20/C-27 peptide, all but one, 45C11, still bind to the PSMGFR peptide, albeit at the N-terminal portion of PSMGFR. The binding of 45C11 is weak but deductive reasoning shows that all or some of the cognate epitope must lie within SNIKFRPGSVV (SEQ ID NO:1744).

**[00634]** Of the antibodies generated by immunizing with the N+9/C-9 peptide, 8A9 and 17H6 do not bind to the PSMGFR peptide, so must bind to the 9 additional N-terminal amino acids. Antibodies 3C5 and 39H5 appear to bind to the 10 most N-terminal amino acids of the PSMGFR peptide.

**[00635]** In order to further refine the epitopes to which each antibody binds, a series of smaller peptides derived from the PSMGFR sequence were synthesized: N-30 (SEQ ID NO:7), N-26 (SEQ ID NO:6), N-19 (SEQ ID NO:4), N-10/C-5 (SEQ ID NO:8), N-19/C-5 (SEQ ID NO:9). Each of the antibodies was tested in an ELISA assay for their ability to bind to this refined set of peptides, plus PSMGFR, N-10 and C-10 peptides (Fig. 67-69).

**[00636]** In **Fig. 67A-67D**, antibodies generated by immunization with the PSMGFR peptide were assayed. As can be seen in the figure, amino acids ASRYNLT (SEQ ID NO:1745), which are essentially in the middle of the PSMGFR peptide, are important or essential for the binding of 28F9, 18G12, 25E6, and MNC3 antibodies. Amino acids GTINVHDTVET (SEQ ID NO:1746), which comprise the most N-terminal part of the PSMGFR peptide are important or essential for the binding of the 18B4 antibody. Amino acids FPFS (SEQ ID NO:1747) are important or essential for the binding of 20A10, MNC2 and MNE6. We note that these three antibodies recognize a conformational epitope, not a linear epitope. Because the proline in the FPFS sequence significantly alters the conformation of nearby portions of the PSMGFR peptide, it is also possible that the antibodies do not bind directly to these four amino acids, but that the absence of the proline alters the fold of the remaining peptide such that the conformation to which 20A10, MNC2 and MNE6 bind, is no longer present.

**[00637]** In **Fig. 68A-68D**, antibodies generated by immunization with the N+20/C-27 peptide were assayed. As can be seen in the figure, amino acids GTINVHDTVET, which comprise the most N-terminal part of the PSMGFR peptide are important or essential for the binding of the 29H1, 32C1, and 31A1 antibodies. Amino acids SNIKFRPGSVVVQLTLAFRE (SEQ ID NO:1748), which is 20 additional amino acids N-terminal to the PSMGFR peptide and outside of the PSMGFR peptide, are important or essential for the binding of antibody 45C11. However,



referring back to **Fig. 65**, antibody 45C11 was not able to bind to the N+9/C-9 peptide, therefore we conclude that amino acids within the SNIKFRPGSVV sequence are essential for the binding of 45C11. Amino acids QFNQYKTEA (SEQ ID NO:1749), which are still within the sequence of PSMGFR, are important or essential for the binding of antibody 1E4.

**[00638]** In **Fig. 69A-69D**, antibodies generated by immunization with the N+9/C-9 peptide were assayed. As can be seen in the figure, amino acids GTINVHDTVET, which comprise the most N-terminal part of the PSMGFR peptide are important or essential for the binding of the 39H5 and 3C5 antibodies. As can be seen in the figure, amino acids VQLTLAFRE (SEQ ID NO:1750), which is 9 additional amino acids N-terminal to the PSMGFR peptide and outside of the PSMGFR peptide, are important or essential for the binding of antibodies 17H6 and 8A9. Because the 17H6 and 8A9 antibodies do not bind to any of the smaller peptides shown in this figure, refer to **Figure 66A-66C**, which shows that these two antibodies only bind to the peptide that has 9 additional amino acids N-terminal to the PSMGFR peptide.

**[00639]** Table 2 below lists antibodies of the invention and their cognate epitopes.

<b>Table 2</b>		
<b>Immunizing Peptide</b>	<b>Antibody Name</b>	<b>Cognate Sequence</b>
PSMGFR	MNC2	FPFS or PFPFSAQSGA
	MNE6	FPFS or PFPFSAQSGA
	20A10	FPFS or PFPFSAQSGA
	3C2B1	FPFS or PFPFSAQSGA
	5C6F3	SVSDV
	MNC3	ASRYNLT
	25E6	ASRYNLT
	28F9	ASRYNLT
	18G12	ASRYNLT
	18B4	GTINVHDTVET
N+20/C-27		
	45C11	SNIKFRPGSVV
	29H1	GTINVHDTVET
	32C1	GTINVHDTVET
	31A1	GTINVHDTVET
	1E4	QFNQYKTEA
N+9/C-9		
	17H6	VQLTLAFRE
	8A9	VQLTLAFRE
	39H5	GTINVHDTVET

	3C5	GTINVHDTVET

**[00640] Ability to displace NME7<sub>AB</sub> binding to the MUC1\* extra cellular domain peptide PSMGFR**

**[00641]** We previously reported that dimeric NME1 dimerizes MUC1\* extra cellular domain and stimulates growth. Monomeric NME7<sub>AB</sub> has two binding sites for MUC1\* so that as a monomer it dimerizes MUC1\* and mediates cancer cell growth. We showed that NME1 and NME7<sub>AB</sub> can bind to the MUC1\* extra cellular domain. In vitro, NME1 and NME7<sub>AB</sub> bind to the PSMGFR peptide even if the 10 N-terminal amino acids are deleted, referred to herein as N-10 (SEQ ID NO:3). However, neither NME1 nor NME7<sub>AB</sub> can bind to the PSMGFR peptide if the 10 membrane proximal amino acids are deleted, referred to herein as C-10 (SEQ ID NO:825). In summary, the epitope to which NME1 and NME7<sub>AB</sub> bind includes all or part of the 10 membrane proximal amino acids: PFPFSAQSGA (SEQ ID NO:1743). We tested various antibodies that were generated in animals by immunizing with the PSMGFR peptide for their ability to recognize cancer cells but not healthy cells. Among the most cancer selective were the MNC2 and MNE6 monoclonal anti-MUC1\* antibodies. Two other monoclonal antibodies that were generated from immunizing animals with the PSMGFR peptide are MNC3 and MNC8. Although MNC2, MNE6, MNC3 and MNC8 all bind to the PSMGFR peptide, like NME1 and NME7<sub>AB</sub>, MNC2 and MNE6 bind strongly to the N-10 peptide but not to the C-10 peptide. In fact, MNC2 and MNE6 competitively inhibit the binding of NME1 and NME7<sub>AB</sub> to PSMGFR. Conversely, MNC3 and MNC8 are able to bind to the C-10 peptide, bind less well to the N-10 peptide and do not compete with NME1 nor NME7<sub>AB</sub> for binding to MUC1\* peptides, including PSMGFR (Fig. 70). MNC3 and MNC8 are less cancer specific than MNC2 and MNE6. MNC3 and MNC8 recognize stem and progenitor cells, such as hematopoietic stem cells, whereas MNC2 and MNE6 do not. Because hematopoietic stem cells are the progenitor cells for the blood cells, it would be problematic to have a cancer therapeutic that would also target such an important normal cell type.

**[00642]** In this experiment, antibodies of the invention were tested for their ability to displace NME7<sub>AB</sub> from binding to the PSMGFR peptide. In this experiment, a multi-well plate was coated with the PSMGFR peptide. Recombinant NME7<sub>AB</sub> was allowed to bind to the surface-

immobilized PSMGFR peptide. Wash steps followed. Various antibodies were added, followed by wash steps. The amount of NME7<sub>AB</sub> that remained attached to the PSMGFR coated plate, after antibody competition, was measured by detecting a tag on the NME7<sub>AB</sub>. As a control, anti-NME7<sub>AB</sub> antibodies were also tested for their ability to displace NME7<sub>AB</sub> from the PSMGFR. Figure 70 shows a graph of an ELISA displacement assay. The bar graph is color coded to indicate the cognate epitope to which each antibody binds. As can be seen in the figure, the antibodies that bind to the more C-terminal portions of PSMGFR are the most potent at disrupting the binding of onco-embryonic growth factor NME7<sub>AB</sub> to the MUC1\* extra cellular domain or the PSMGFR peptide. The rank order of potency for disrupting binding of NME7<sub>AB</sub> to PSMGFR according to their cognate epitope is as follows: FPFS> ASRYNLT> QFNQYKTEA>GTINVHDTVET. Antibodies that bind to epitopes outside of the PSMGFR peptide, such as 45C11, 8A9 and 17H6 did not compete with NME7<sub>AB</sub> for binding.

**[00643] Western blot assay to determine linear versus conformational cognate epitope**

**[00644]** Antibodies were tested to determine whether they recognize a linear or a conformational epitope. Only antibodies that recognize a linear epitope work in Western blots when using denaturing gels. For comparison, known antibodies were tested for their ability to bind to HCT-116, a MUC1 negative cancer cell line, HCT-MUC1-18, which is a cleavage resistant clone of HCTs transfected with full-length MUC1, and HCTs transfected with MUC1\*, wherein the extra cellular domain comprises only the PSMGFR sequence. The antibodies tested for comparison are MNC2 and MNE6, which were known to only recognize a conformational epitope, SDIX which is a polyclonal antibody raised against PSMGFR and VU4H5, which is a commercially available monoclonal antibody that recognizes the tandem repeats of full-length MUC1 (**Fig. 71A-71D**). As can be seen, neither MNC2 nor MNE6 recognize a MUC1 or MUC1\* specific linear epitope. The SDIX polyclonal antibody recognizes HCT-MUC1\* but not full-length MUC1 and VU4H5 only recognizes full-length MUC1. These same antibodies were also tested for their ability to work in Western blots of two breast cancer cell lines 1500, aka Zr-75-1, and T47D cells and show the same binding pattern (**Fig. 71E-71H**).

**[00645]** Antibodies that were raised against the PSMGFR peptide were tested the same way in Western blots (**Fig. 72A-72P**). As can be seen, antibodies 25E6 and 18B4 recognize linear epitopes but 20A10, 3C2B1, 5C6F3, 18G12 and 28F9 do not, indicating that they bind to a conformational epitope. Antibodies that were raised against the N+20/C-27 peptide were tested

the same way in Western blots (**Fig. 73A-73J**). As can be seen, antibodies 31A1 and 32C1 recognize linear epitopes. Antibodies 1E4 and 45C11 may recognize a conformational epitope. Antibodies that were raised against the N+9/C-9 peptide were tested the same way in Western blots (**Fig. 74A-74H**). As can be seen, none of these antibodies recognize linear MUC1 or MUC1\* specific epitopes. These antibodies may recognize a conformational epitope. However, an alternative interpretation is that the lack of binding in a Western blot means that they do not specifically recognize MUC1 or a MUC1 cleavage product or that the concentration used in this assay was insufficient.

**[00646] Recognition of a MUC1 cleavage product after cleavage by MMP9**

**[00647]** We previously demonstrated that MNC2 recognizes a MUC1\* that is generated when full-length MUC1 is cleaved by matrix metalloprotease 9, MMP9 (**Fig. 37**). MMP9 is expressed by tumor tissues and is a predictor of poor prognosis for breast cancers (vant Veer et al 2002; Dufour et al 2011). MMP9 has also been implicated in metastasis (Owyong et al 2019). Recall also that MNC2 competitively inhibits the binding of onco-embryonic growth factor NME7<sub>AB</sub> to the MUC1\* extra cellular domain (**Fig. 3**). Therefore, it follows that onco-embryonic growth factor, which activates growth and survival functions of MUC1\*, also recognizes a MUC1\* generated by cleavage by MMP9. It then follows that the most cancer specific antibodies are those that recognize a conformational epitope formed when MUC1 is cleaved to MUC1\* by MMP9.

**[00648]** Antibodies generated by immunization with PSMGFR, N+20/C-27, or N+9/C-9 were tested for their ability to recognize MUC1 after it is cleaved by MMP9. To do this, we transfected HCT-116, a MUC1 negative colon cancer cell line, with full-length MUC1 and isolated a single cell clone that is cleavage resistant; this cleavage resistant cell line is called HCT-MUC1-18. To HCT-MUC1-18 cells was added either a catalytically active MMP9 or MMP2. The enzymes, added over a range of concentrations, were incubated with the cells for 24 hours. The resultant cells were then incubated with the various antibodies and analyzed by FACS to determine which bound to a MUC1 cleavage product produced by cleavage by MMP9 (**Fig. 75A-75N**). Note that the first bar of each graph shows that none of the antibodies binds to full-length MUC1 in the absence of cleavage. Each bar graph is labeled with both the name of the antibody used in that assay and its cognate epitope. The order of the graphs from right to left corresponds to the distance from the cell surface of the antibody's cognate epitope. The

antibodies that bind to the more C-terminal epitopes within PSMGFR peptide, such as 20A10 and 25E6, showed the most increased binding to a MUC1 cleavage product after cleavage by MMP9 but not MMP2. Antibody 45C11, which binds to the SNIKFRPGSVV epitope, which is outside of the PSMGFR portion of MUC1, does not recognize a MUC1 cleavage product after cleavage by MMP9 or MMP2 (Fig. 75K). Similarly, antibodies 8A9 and 17H6 bind to the VQLTLAFRE epitope, which is also outside of the PSMGFR sequence, and they do not bind to a MUC1 cleaved by MMP9 or MMP2. This result is consistent with the idea that MMP9 cleaves MUC1 such that the extra cellular domain of the remaining transmembrane cleavage product comprises essentially the amino acids of the PSMGFR peptide. For the greatest degree of cancer specificity, the antibody should recognize a conformational epitope of a MUC1 cleavage product created when MUC1 is cleaved by MMP9. Of the antibodies shown in Fig. 75A-75N, only 20A10 recognizes the MUC1 cleavage product produced by cleavage by MMP9 and also does not work in a Western blot, indicating it recognizes a conformational epitope, as do MNC2 and MNE6. Cleavage and release of the massive tandem repeat domain of MUC1 unmask the ectopic binding site on MUC1\*; linear epitopes will be unmasked in addition to conformational epitopes.

**[00649] FACS analysis of binding to a panel of cancer cell lines**

**[00650]** Fluorescence Activated Cell Sorting, FACS, was performed on reference antibodies as well as new antibodies of the invention. FACS analyses of reference antibodies MNC2, “C2”, and VU4H5 binding to either the MUC1-negative cell line HCT-116, HCTs transfected with MUC1\*, “HCT-MUC1\*”, a cleavage resistant single cell clone of HCTs transfected with MUC1 full-length, “HCT-MUC1-18”, and MNC2 binding to breast cancer cells line T47D or breast cancer cell line 1500 also known as ZR-75-1, was performed (Fig. 76A-76J). This analysis shows that MNC2 binds to an ectopic binding site on the extra cellular domain of MUC1\*, which is only available after cleavage and release of the bulk of the extra cellular domain comprising the tandem repeat domain. VU4H5 binds to hundreds of repeating epitopes in the tandem repeat domain of full-length MUC1 and does not bind to MUC1\*. Although we know that cancer cell lines express both full-length MUC1 and MUC1\*, antibodies against full-length MUC1 have, as yet, been shown to have no therapeutic value. Stimuvax, ImMucin, IMG242, SAR566658, PankoMab and AS1402 were all antibodies that bound to full-length MUC1 and all failed to show efficacy in clinical trials. MUC1\*, and not full-length MUC1, is a potent growth

factor receptor that mediates the growth of cancer cells (Mahanta et al 2008) and their resistance to chemotherapy agents (Fessler et al 2009). These studies showed that full-length MUC1 had no tumor promoting activity. Further, IHC studies show that as tumor stage increases, the amount of MUC1\* increases as the amount of full-length MUC1 decreases (Fig. 54). In fact, studies with tissue micro arrays of breast cancers show that nearly 30% of breast cancer specimens had no detectable full-length MUC1, compared to only 5% that were negative for MUC1\* (Fig. 10-11). A point to consider for therapeutics that target full-length MUC1 is that if cells expressing full-length MUC1 are eliminated, that would simply enrich the tumor population for the more virulent MUC1\* growth factor receptor expressing cells, which would make the cancers worse.

**[00651]** Reference antibody MNC2, “C2”, was analyzed by FACS for its ability to bind to a panel of cancer cell lines that are all MUC1\* positive, with the exception of MDA-MB-231, which expresses MUC1 and MUC1\* at a level that is so low that it is often used as a negative control (Fig. 77A-77N). The panel of cancer cells that was probed with MNC2 included T47D and 1500 breast cancer cells, NCI-H292 and NCI-H1975 lung cancer cells, SKOV-3 ovarian cancer cells, HPAF-II and Capan-1 pancreatic cancer cells, DU145 prostate cancer cells, and MDA-MB-231, breast cancer cells, which are nearly MUC1 and MUC1\* negative. MNC2 robustly recognized a wide range of cancer cell lines. We note that although MNC2 recognized HPAF-II pancreatic cells, it did not recognize another pancreatic cell line, Capan-1, as well. Similarly, MNC2 did not recognize prostate cancer cell line DU145 very well. In IHC tissue studies, we found that MNC2 recognized about 57% of prostate cancer tissues and 78% of pancreatic tissues, albeit with significant tumor heterogeneity.

**[00652]** **Figure 78A-78C** shows a color coded schematic of the PSMGFR sequence that has been extended or deleted at both the N- and C-termini. Antibodies of the invention were tested against this subset of peptides to further refine the epitopes to which each antibody binds or the critical amino acids within the epitope to which each antibody binds. Fig. 78A is an aligned schematic of the various subsets of peptides. Fig. 78B lists the antibodies that bind to each of the color coded sequences. Fig. 78C lists the cancer cell lines that each antibody recognizes.

**[00653]** Figures 80-87 show graphs of FACS analyses wherein antibodies of the invention are compared for their ability to specifically recognize different types of cancer cells. Percent cells recognized as well as the Mean Fluorescence Intensity, MFI, was measured. Considering only these FACS experiments, they show that only antibodies that recognize the PSMGFR peptide are

able to recognize cancer cell lines. Antibodies that bind to epitopes outside of the PSMGFR sequence do not specifically recognize these cancer cell lines.

**[00654] IHC tissue studies of normal versus cancerous tissues to determine true cancer specificity**

**[00655]** Immunohistochemistry, IHC, tissue studies of tissue micro arrays, “TMAs”, are a more stringent test of the cancer specificity of antibodies than FACS analysis of a single cancer cell line. Cancer cell lines are a single cell from a single patient that have been expanded in a lab for decades. Cell lines are limited in that they are not representative of a cross section of the human population. Further, after culturing the cell line in vitro for decades it may no longer look like the original cell. Also, there are no real normal cell lines for comparison, as they have to be made immortal. Tissue studies are more informative because each tissue micro array comprises tissues from multiple donors and the cells are in their natural environment, without years of culturing under non-physiologic conditions. Additionally, tissues provide information regarding tumor heterogeneity as well as information regarding normal patterns of expression. Each antibody of the invention was used to probe a normal tissue micro array, FDA Normal Array 1021. In addition the antibody was also used to probe a panel of cancerous tissue arrays. In some cases, antibodies that showed strong staining of normal tissues, especially of critical organs such as heart or lung, were tested on a limited number of cancerous tissue arrays, since their cross reactivity to normal tissues eliminated them from consideration as anti-cancer therapeutics.

**[00656] Figures 113-200** show photographs of the IHC staining of normal TMAs versus cancerous TMAs for each antibody of the invention.

**[00657] Figure 113-120** show photographs of tissues studies probed with antibody 20A10. Recall that 20A10 binds to the PSMGFR peptide, binds to the N-10 peptide, but does not bind to the C-10 peptide. Refined epitope mapping shows that like MNC2 and MNE6, the binding of 20A10 depends on amino acids FPFS being present in the PSMGFR peptide. 20A10 binds to the most membrane proximal part of the MUC1\* extra cellular domain. An overview of FDA Normal Tissue Array 1021 is shown in Figure 113. Figure 114A-114X show that there is little to no cross reactivity of 20A10 for normal tissues. We note that MNC2, MNE6 and 20A10 all react with the MUC1\* that is expressed on the luminal edge of the terminal breast ducts, luminal edge of the fallopian tubes, luminal edge of about 10% of the distal collecting ducts of normal kidney, and luminal edge of ureter. Because the staining is strictly limited to the luminal edge of a subset

of ducts and glands, these antibodies are considered to be safe as therapeutics as the inside of ducts and glands are protected from large entities carried by blood, such as antibodies or CAR T cells. Importantly, MNC2, MNE6 and 20A10 show no staining of critical organs, such as heart, lung and brain. In stark contrast, 20A10, like MNC2 and MNE6, robustly binds to cancerous tissues. 20A10 stains nearly all specimens of the BR1141 breast cancer array (Fig. 115-116). In addition to robust staining of the breast cancer tissue, the staining is membrane staining, indicating that 20A10 recognizes an extra cellular portion of MUC1\*, which is critical for an effective antibody-based anti-cancer therapeutic. 20A10 also showed robust and membranous staining of pancreatic cancer tissues (Fig. 117-118) and esophageal cancer tissues (Fig. 119-120). In summary, 20A10 shows great cancer specificity and as an anti-cancer therapeutic offers a large therapeutic window because of the vast difference between staining of normal tissues and cancerous tissues, in terms of the location and intensity of staining.

**[00658]** Anti-MUC1\* antibody 3C2B1 is an antibody that like MNC2, MNE6 and 20A10, binds to N-10 but not to C-10. More refined epitope mapping shows that like these three other highly cancer-specific antibodies, 3C2B1 requires the FPFS sequence for binding to a MUC1\* extra cellular domain peptide. **Figure 121** shows the photograph of the FDA normal array 1021. **Figure 122A-122X** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL. As can be seen, there is no binding of 3C2B1 to any critical normal organs. **Figure 123** shows photograph of pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 3C2B1 at 1-20ug/mL. **Figure 124** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL. **Figure 125** shows photograph of breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL. **Figure 126A-126F** shows magnified photographs of specific tissues from breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 3C2B1 at 20ug/mL. As can be seen in the figure, 3C2B1 robustly stains breast cancer tissues.

**[00659]** Anti-MUC1\* antibody 5C6F3 binds to the N-10 peptide, does bind to the C-10 peptide, although binding is reduced somewhat. Its cognate epitope comprises all or some of the sequence SVSDV (SEQ ID NO:1751). **Figure 127** shows photograph of FDA normal tissue array 1021 stained with the anti-PSMGFR antibody 5C6F3 at 1ug/mL. **Figure 128** shows photographs of specific tissues from FDA normal tissue array 1021 stained with the anti-



PSMGFR antibody 5C6F3 at 1ug/mL. **Figure 129** shows photograph of pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 5C6F3 at 1-20ug/mL. **Figure 130** shows photographs of specific tissues from pancreatic cancer tissue array PA1003 stained with the anti-PSMGFR antibody 5C6F3 at 1ug/mL. **Figure 131** shows photograph of breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 5C6F3 at 1ug/mL. **Figure 132** shows photographs of specific tissues from breast cancer tissue array 1141 stained with the anti-PSMGFR antibody 5C6F3 at 1ug/mL. As can be seen in the figure 5C6F3 is a high affinity antibody that has great cancer-specificity and with the exception of adrenal, which may be an artefact of that tissue, did not show binding to normal tissues.

**[00660]** In contrast to 20A10, which binds to the most membrane proximal part of the MUC1\* extra cellular domain, **18B4** binds within the GTINVHDTVET sequence, which is the most distal part of the PSMGFR sequence. Unlike antibodies MNC2, MNE6 or 20A10, 18B4 cannot bind to the N-10 peptide but does bind to the C-10 peptide. Figure 133-134 show the binding of antibody 18B4 to normal tissues. In contrast to 20A10, antibody 18B4 shows strong binding to a wide range of normal tissues (Fig. 134), including lung (Fig. 134K). Figure 135-138 show 18B4 staining of breast cancer tissues and esophageal cancer tissues. Because of the strong binding of 18B4 to normal tissues, there is less cancer specificity to this antibody.

**[00661]** **Figure 139-144** show the binding of PSMGFR antibody **18G12** to normal tissues, breast cancer tissues and esophageal cancer tissues. 18G12 is able to bind to the N-10 peptide, but is also able to bind to the C-10 peptide. 18G12 binds to the ASRYNLT epitope within the PSMGFR peptide. Antibody 18G12 binds to the luminal edge of many of the collecting ducts of normal kidney (Fig. 140D), binds to normal heart muscle (Fig. 140I) as well as to normal skeletal muscle (Fig. 140X). However, there is a clear cancer specificity in that 18G12 binds much more strongly to cancerous tissues than to the few normal tissues. In addition, 18G12 stains the entire cancerous tissues rather than just a luminal edge here or there. Figure 141-146 show 18G12 staining of breast cancer tissues, pancreatic cancer tissues and esophageal cancerous tissues. The contrast between the staining of the normal tissues and the cancer tissues clearly demonstrates cancer specificity.

**[00662]** **Figure 147-148** show the binding of PSMGFR antibody 25E6 to normal tissues. 25E6 is able to bind to the N-10 peptide, but is also able to bind to the C-10 peptide. 25E6 binds to the ASRYNLT epitope within the PSMGFR peptide. Like MNC2, MNE6 and 20A10,

antibody 25E6 binds to the luminal edge of terminal breast ducts, luminal edge of fallopian tubes, to the luminal edge of a subset of the distal collecting ducts of normal kidney and to the luminal edge of ureter. Unlike MNC2, MNE6 and 20A10, 25E6 binds, albeit very weakly, to normal heart muscle (Fig. 148I) as well as to normal skeletal muscle (Fig. 148X). However, there is a clear cancer specificity in that 25E6 binds much more strongly to cancerous tissues than to the few normal tissues. In addition, 25E6 stains the entire cancerous tissues rather than just a luminal edge here or there. Figure 149-152 show 25E6 staining of breast cancer tissues and pancreatic cancerous tissues. The contrast between the staining of the normal tissues and the cancer tissues clearly demonstrates cancer specificity.

**[00663]** Figure 153-156 show the binding of PSMGFR antibody **28F9** to normal tissues and breast cancer tissues. 28F9 is able to bind to the N-10 peptide, but is also able to bind to the C-10 peptide. 28F9 binds to the ASRYNLT epitope within the PSMGFR peptide. Like MNC2, MNE6 and 20A10, antibody 25E6 binds to the luminal edge of terminal breast ducts, luminal edge of fallopian tubes, to the luminal edge of a subset of the distal collecting ducts of normal kidney and to the luminal edge of ureter. Figure 155-156 show 28F9 staining of breast cancer tissues..

**[00664]** Figure 157-158 show the binding of the N+20/C-27 antibody **1E4** to normal tissues. 1E4 is able to bind to the N-10 peptide but also is able to bind to the C-10 peptide. 1E4 binds to the QFNQYKTEA sequence which is within the PSMGFR sequence. Examination of the entire normal tissue micro array (Fig. 157A) shows that antibody 1E4 binds to many normal tissues, including brain, cerebellum, all 3 liver specimens, pancreas, parathyroid, spinal cord and skeletal muscle. Magnified images show that 1E4 stains heart (Fig. 158I) as well. 1E4 staining of a breast cancer array (Fig. 159-160) shows that there is some cancer specificity.

**[00665]** Figure 161-162 show the binding of the N+20/C-27 antibody **29H1** to normal tissues. **29H1** binds within the GTINVHDVET sequence, which is the most distal part of the PSMGFR sequence. Unlike antibodies MNC2, MNE6 or 20A10, 29H1 cannot bind to the N-10 peptide but does bind to the C-10 peptide. Examination of the entire normal tissue micro array (Fig. 157A) shows that even at concentration as low as 0.5 ug/mL, antibody 29H1 strongly stains a wide range of normal tissues, including brain, heart, liver and lung. 29H1 staining of a breast cancer array (Fig. 163-164) and staining of a pancreatic cancer tissue array (Fig. 165-166) shows that there is no cancer specificity.

**[00666]** Antibody 31A1 is similar to 29H1 in that they are both N+20/C-27 antibodies that bind within the GTINVHDTVET (SEQ ID NO:1746) sequence, which is the most distal part of the PSMGFR sequence. Unlike antibodies MNC2, MNE6 or 20A10, neither 31A1 nor 29H1 can bind to the N-10 peptide but do bind to the C-10 peptide. Examination of the entire normal tissue micro array and the magnified images (Fig. 167-168) shows that even at concentration as low as 0.5 ug/mL, antibody 31A1 strongly stains a wide range of normal tissues, including brain, heart, lung, spleen, bone marrow, and skeletal muscle. 31A1 was used to stain a breast cancer array, (Fig. 169-170). 31A1 was used over a range of concentrations to stain a pancreatic cancer tissue array (Fig. 171-172). These figure shows that 31A1 has insufficient cancer specificity.

**[00667]** Antibody **32C1** is similar to 29H1 and 31A1 in that they are all N+20/C-27 antibodies that bind within the GTINVHDTVET sequence, which is the most distal part of the PSMGFR sequence. Unlike antibodies MNC2, MNE6 or 20A10, none of 32C1, 31A1 or 29H1 can bind to the N-10 peptide but all do bind to the C-10 peptide. Examination of the entire normal tissue micro array and the magnified images (Fig. 173-174) shows that even at concentration as low as 0.25 ug/mL, antibody 32C1 strongly stains a wide range of normal tissues, including brain, heart, lung, liver, spleen and bone marrow. 32C1 was also used to probe a breast cancer array (Fig. 175-176). 32C1 was used over a range of concentrations to stain an esophageal cancer tissue array (Fig. 177-178). Taken together, these figures show that 32C1 has insufficient cancer specificity.

**[00668]** Antibody **45C11** is an N+20/C-27 antibody that binds to epitope SNIKFRPGSVV (SEQ ID NO:1744) that is 20 amino acids outside of the PSMGFR sequence at the N-terminal end. 45C11 does not bind to the N-10 peptide. Normal tissue array FDA 1021 was stained with 45C11 at 12.5ug/mL (Fig. 179-180). As can be seen in the figures, 45C11 shows strong binding to many normal tissues, including brain, heart, lung, liver, spleen, skeletal muscle and bone marrow. 45C11 was used over a range of concentrations to stain a breast cancer tissue array (Fig. 181-182). 45C11 was also used to stain a pancreatic cancer tissue array (Fig. 183-184). Taken together, these figures show that 45C11 has no cancer specificity.

**[00669]** Antibody **3C5** is an N+9/C-9 antibody that binds to epitope GTINVHDTVET. Like the other antibodies that bind to this epitope such as 32C1, 29H1 and 31A1, they bind to the most distal, that is to say the most N-terminal, part of the PSMGFR sequence. Unlike antibodies MNC2, MNE6 or 20A10, none of 3C5, 32C1, 31A1 or 29H1 can bind to the N-10 peptide but all

do bind to the C-10 peptide. Examination of the entire normal tissue micro array, where 3C5 was used at 10ug/mL, and the magnified images (Fig. 185-186) shows that antibody 3C5 strongly stains some normal tissues, including brain, heart, adrenal gland and bone marrow. 3C5 was also used to probe a pancreatic cancer array at 10ug/mL, (Fig. 187-188). Taken together, these figures show that 3C5 has no cancer specificity.

**[00670]** Antibody **8A9** is an N+9/C-9 antibody that binds to epitope VQLTLAFRE which is outside of the PSMGFR sequence. Antibody 8A9 cannot bind to the N-10 peptide. Normal tissue array FDA 1021 was stained with 8A9 (Fig. 189-190). As can be seen in the figures, like antibody 45C11, which also binds an epitope that is N-terminal beyond the PSMGFR sequence, antibody 8A9 shows strong binding to many normal tissues, including adrenal, brain, heart, lung, liver, spleen, skeletal muscle and bone marrow. A pancreatic cancer array stained with antibody 8A9 showed weak binding to a subset of pancreatic cancer tissues (Fig. 191-192). Taken together, these figures show that 8A9 has no cancer specificity.

**[00671]** Antibody **17H6** is an N+9/C-9 antibody that binds to epitope VQLTLAFRE, which is outside of the PSMGFR sequence. 17H6 was used to stain normal tissue array 1021. Examination of the entire normal tissue micro array and the magnified images (Fig. 193-194) shows that antibody 17H6 stains some normal tissues, including brain, heart, adrenal gland, bone marrow and skeletal muscle. 17H6 was used to probe a pancreatic cancer array and showed weak binding to most pancreatic cancer tissues (Fig. 195-196). However, the binding of 17H6 to several normal tissues of critical organs shows that 17H6 has little cancer specificity.

**[00672]** Antibody **39H5** is an N+9/C-9 antibody that binds weakly to the intact PSMGFR peptide but not significantly to any of the subset peptides. 39H5 may bind to the GTINVHDTVET, which is the most distal part of the PSMGFR sequence. Examination of the entire normal tissue micro array and the magnified images (Fig. 197-198) shows that antibody 39H5 stains some normal tissues, including brain, heart, liver and bone marrow. 39H5 was used to probe a pancreatic cancer array, (Fig. 199-200). Although 39H5 stained a good percentage of the pancreatic cancer specimens, considering the normal tissues that 39H5 stained, 39H5 has little cancer specificity.

**[00673] Summary of FACS analysis**

**[00674]** Determining the cancer specificity of antibodies using cell lines is difficult, as these cells were obtained from a single patient's tumor decades ago, and then propagated in culture for

decades. Even if the patient's tumor was at one point heterogeneous, the decades of in vitro culture have essentially made the cell line a single cell clone. Antibodies of the invention were assayed by FACS to determine if they bound to MUC1 or MUC1\* positive cancer cells but not MUC1 negative cells. The results of these experiments are shown in Figures 76-87. What is very clear is that antibodies that bind to epitopes of the MUC1 sequence that are outside of and N-terminal to PSMGFR sequence show no cancer specificity. Referring now to the readings of Mean Fluorescence Intensity (MFI) it appears that antibodies with cognate epitopes at the very N-terminus of the PSMGFR sequence, such as those that bind to an epitope within GTINVHDTVET, show far less cancer specificity than the antibodies that recognize more C-terminal epitopes. For example, antibody MNC2 that will not bind to the C-10 peptide binds strongly to nearly every MUC1\* positive cell line (Fig. 76-77). However, closer examination reveals that MNC2 binds lung cancer line NCI-H1975 much more strongly than NCI-H292. Similarly, MNC2 binds pancreatic cell line HPAF-II much better than Capan-1 or prostate cancer line DU145. PCR measurements show that the expression levels of cleavage enzymes varies greatly across a panel of cancer cell lines (Fig. 43 and Fig. 44 ). The fold of the MUC1\* extra cellular domain can vary greatly depending on which cleavage enzyme clips it, which likely accounts for differences between cancer cell lines that a single antibody recognizes. This variation in antibody recognition of various cell lines, even within a cancer sub-type is apparent in the figures.

**[00675] Summary of IHC data**

**[00676]** IHC analysis of real tissues, including both normal and cancerous tissues, is more informative than the study of cultured cell lines, as is necessary in FACS analysis. Each antibody was first tested over a range of concentrations to determine optimal concentration. Antibody concentration was increased until the stroma also picked up stain, which indicates non-specific background binding. The optimal concentration for that particular antibody was then deemed to be just below the concentration at which the antibody stained the stroma.

**[00677]** An overview of the IHC tissue studies is shown in Figure 88-112. Here, we focused on the binding of antibodies to critical organ tissues, since binding to certain normal tissues would likely eliminate therapeutic use of that antibody. In these figures, the antibodies were grouped according to their cognate epitope. What is evident from the tissue studies is that the further the epitope is from the cell membrane, the more it binds to normal MUC1 on normal

tissues. For example, binding to normal heart tissue by representative antibodies that recognize a specific epitope are shown in Figure 88A-88L. As the figure illustrates, antibodies that bind to epitopes that are N-terminal to the PSMGFR peptide such as epitope within SNIKFRPGSVV or VQLTLAFRE show such strong binding to normal heart that they could not be used in therapeutics. In addition, antibodies that bind to the more N-terminal portion of PSMGFR, such as 29H1, also show binding to normal heart. The antibodies with the least binding to normal tissues and the strongest binding to cancerous tissues bind to epitopes within the FPFS or PFPFSAQSGA. Some antibodies that bind to epitopes within the ASRYNLT portion may also be suitable as therapeutics. These antibodies and others that recognize the same epitopes are desirable as anti-cancer therapeutics because they have a large therapeutic window, meaning that because of the low binding to normal tissues, and low side effects, patients can be dosed with antibody levels high enough to effectively kill the tumor cells. More detailed photographs of antibodies of the invention binding, or not binding, to other critical tissues are also shown. Figures 89-94 show magnified photographs of each antibody binding to normal heart tissue, where the antibodies have been categorized according to which epitope they bind. Figures 95-100 show magnified photographs of each antibody binding to normal liver tissue, where the antibodies have been categorized according to which epitope they bind. Figures 101-106 show magnified photographs of each antibody binding to normal lung tissue, where the antibodies have been categorized according to which epitope they bind. Figures 107-112 show magnified photographs of each antibody binding to normal bone marrow, where the antibodies have been categorized according to which epitope they bind.

[00678] The results of the IHC studies (Fig. 88-Fig. 200) are summarized in **Table 3**.

**Table 3: Summary of Antibody Cross-Reactivity to Normal Tissues**

Antibody Name	HEART	BRAIN	LUNG	LIVER	SPLEEN	BONE MARROW	KIDNEY	SKELETAL MUSCLE	ADRENAL
PSMGFR									
MNC2									
MNE6									
20A10									
25E6	~+							~+	

18B4			+++			+			
18G12	++						+++	+++	
28F9								~+	
3C2B1									
5C6F3			+			~+	++		++++
<b>N+20/C-27</b>									
1E4	++	++		+++		~+		+++	
31A1	++	+++	++		++++	++++		+	
32C1	+++	+++	+++	++	+++	++++			++
29H1	+	++++	++++	++++	++				++++
45C11	+++++	++++	++	+++++	++	+++	+	+++++	+
<b>N+9/C-9</b>									
8A9	+++++	+++++	+++	+++++	++++	++++	+	++++	+++
17H6	+++	++++				+			++
3C5	+++	++++		+		++++			+++
39H5	++++	+++++	+	+++	+++	++++			++++

**[00679]** As can be clearly seen in the table, the further away from the cell membrane that the antibody binds, the more non-specific binding there is. Although these antibodies were generated by immunizing with the PSMGFR peptide, N+20/C-27 peptide or the C+9/C-9 peptide, some of the antibodies generated by immunizing with an extended peptide still bind within the PSMGFR sequence, see Figures 63-69 for the details of epitope binding for each antibody. Some binding to normal tissues can be tolerated if the antibody is incorporated into an appropriate therapeutic format. For example, cellular therapies, such as CAR T, are carried by the blood and meet with physiological barriers including lamina propria and blood-brain barrier that limits the cell's access to luminal edge of ducts and glands. Other antibodies that bind much more strongly to cancerous tissues but do show some binding to normal tissues could also be safe and useful therapeutics if administered locally or if cancer-specificity is enhanced by incorporating into a bi-specific antibody. However, widespread antibody binding to many normal organs or to essential organs for which there is no physical barrier could be lethal to the patient.

**[00680]** The most cancer-specific antibodies with little to no binding to normal tissues are MNC2, MNE6, 20A10, 3C2B1 and 25E6. An ideal antibody therapeutic is one that stains no normal tissues but robustly stains cancer cells. Unfortunately, cancer antigens are also expressed

on normal tissues, so zero staining of normal tissue is not possible. The aim is to identify an antibody that binds much more strongly to tumor tissue than normal tissue and that either binds to non-critical normal tissues or binds to them in a way that would not be physiologically possible in an intact organ. For example, CAR T cells are carried by the blood and the lamina propria is a barrier to their getting to the luminal edge of a duct or gland. Similarly, the blood brain barrier prevents the passage of large molecules like antibodies from the blood into the brain. The usefulness of an antibody as a therapeutic also depends on the format of the therapeutic. As mentioned, cell based therapies have natural barriers that prevent the CAR T cells from getting to some normal tissues. Antibody Drug Conjugate (ADC) based therapies sometimes depend on a local, cancer-specific molecule to activate the toxin attached to the antibody, minimizing the importance of whether or not a naked antibody binds to some normal tissue. In another example, antibodies and antibody-based therapeutics can be administered locally, including intraperitoneally, to maximize the effect on tumor cells while minimizing their effect on normal tissues. In yet another example, an antibody that is not purely cancer-specific can be made more cancer-specific if it is incorporated into a bi-specific antibody where a first side of the molecule binds to a first cancer antigen and the second side of the molecule binds to a second antigen that may be a tissue specific antigen, another cancer specific antigen or even an antigen on a cell such as a T cell, which are called BiTES, bispecific T cell engagers. In yet another example, the less cancer-specific antibody can be incorporated into a cell-based therapy where its expression is induced only after the cell recognizes a tumor. In one aspect, a CAR T cell can express a first CAR that recognizes a first antigen which recognition induces expression of a second antibody, or CAR incorporating the second antibody. In one aspect the cell expresses a CAR directed by an antibody fragment that is cancer-specific and a second antibody or CAR expressing the second antibody is induced to be expressed in an NFAT inducible system. In one aspect the nucleic acids encoding the second antibody or second CAR are down stream of NFAT response elements. The NFAT inducible gene may be inserted into a Foxp3 enhancer or promoter.

**[00681]** Figure 202 shows photographs of pancreatic cancer tissues, each from a different patient. As can be seen, the staining pattern of 1E4 is very different from that of 18B4 and the polyclonal antibody SDIX. 18B4 and SDIX antibodies were generated by immunizing animals with the same peptide (PSMGFR), while the 1E4 antibody was generated from immunization



with a different peptide (N+20). Figures 203-207 show magnified images of selected tissues from this array to highlight the differences between these antibodies. Figure 208 compares the staining of polyclonal antibody SDIX to monoclonal antibody 20A10, which were both generated from immunization with the PSMGFR peptide. Also shown is the difference in staining pattern for antibody 29H1 which was generated by immunization with an N+20 peptide. Although the antibody staining is lighter, antibody 29H1 recognizes more pancreatic cancer tissue specimens than the SDIX polyclonal or 20A10. Figure 209 shows that esophageal cancers are better recognized by antibodies that bind to a MUC1\* peptide with an extended N-terminus, such as antibody 29H1 and antibody 31A1. Similarly, Figure 210 shows that prostate cancers are better recognized by antibodies that bind to a MUC1\* peptide with an extended N-terminus, such as antibody 29H1.

[00682] Below Table 4 shows a summary of the test criteria to determine the cancer-specificity of the various monoclonal antibodies.

**Table 4: Cancer-Specificity Test Criteria**

		1	2	3	4	5	6	7
mAb Name	Binds PSMGFR	Binds N-10	Does not bind C-10	Displaces NME7 <sub>AB</sub> from MUC1*	Does not recognize linear epitope	Recognizes MUC1 after cleavage by MMP9	Cancer selective by FACS	Cancer selective by IHC
VQLTLAFREGTINVHVDVETQFNQYKTEAASRYNLTISDVSVSDVPEPFS AQSGA								
EPFS								
MNC2	☑	☑	☑	☑	☑	☑	☑	☑☑☑
MNE6	☑	☑	☑	☑	☑	☑	☑	☑☑☑
20A10	☑	☑	☑	☑	☑	☑	☑	☑☑☑
3C2B1	☑	☑	☑	☑	☑	☑	☑	☑☑☑
SVSDV								
5C6F3	☑	☑	~☑	☑	☑	☑	☑	☑☑
ASRYNLT								
25E6	☑	☑	☑	☑	☑	☑	☑	☑☑
MNC3	☑	~√	☑	☑	☑	☑	ND	☑
18G12	☑	☑	☑	☑	☑	☑	~√	☑
28F9	☑	☑	☑	☑	☑	☑	~√	☑
QFNQYKTEA								
1E4	☑	☑	☑	☑	☑	☑	~√	☑
GTINVHVDVET								
18B4	☑	☑	☑	~√	☑	☑	☑	☑

29H1	☑	☒	☒	~√	☒	☑	☒	☒
31A1	☑	☒	☒	☑	☒	☑	☒	☒
32C1	☑	☒	☒	~√	☒	☑	☒	☒
39H5	☑	☒	☑	~√	☒	☒	☒	☒
3C5	☑	☒	☑	☒	☒	☒	☒	☒☒
VQLTLAFRE								
8A9	☒	☒	☑	☒	☒	☒	☒	☒☒
17H6	☒	☒	☑	☒	☒	☒	☒	☒
SNIKFRPGSVV								
45C11	☒	☒	☑	☒	☑	☒	☒	☒☒☒

**[00683]** To summarize, we found that antibodies that bound to sequences that are N-terminal to the PSMGFR sequence had no cancer-specificity. Further, the closer to the cell membrane that the antibody binds, the more cancer-specific is the antibody. More importantly, test criteria 1-4 or even 1-5 provide a set of rapid, multiplexed and inexpensive tests that can be performed on hundreds or thousands of impure hybridoma clone supernatants to identify antibodies that are highly selective for cancer-specific forms of MUC1\*.

**[00684] Satisfies test criteria**

**[00685]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on satisfying four (4) of the seven (7) criteria set out in Table 4. In a more preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on satisfying five (5) of the seven (7) criteria set out in Table 4. In a yet more preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on satisfying six (6) of the seven (7) criteria set out in Table 4. In a more preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on satisfying all seven (7) of the criteria set out in Table 4.

**[00686] Bind to N-10**

**[00687]** We have demonstrated that a MUC1 transmembrane protein, devoid of tandem repeats and having an extra cellular domain of 45 amino acids of PSMGFR sequence, is sufficient to function as a growth factor receptor and confers oncogenic characteristics to the cell (Mahanta et al 2008). Antibodies that bind to the PSMGFR peptide or portion of a transmembrane MUC1 cleavage product can be cancer specific but may also bind to stem or progenitor cells. Antibodies that bind to the N-10 peptide are more cancer-specific. In a preferred

embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to bind to the N-10 peptide.

**[00688] Do not bind to C-10**

**[00689]** We have demonstrated that the MUC1 extra cellular domain contains an ectopic binding site that is only exposed if the tandem repeat domain is missing, which can occur as a consequence of alternative splice variant or cleavage and release of the extra cellular domain. Cancer-specific antibodies MNC2 and MNE6 will not bind to full-length MUC1, but do bind to the remaining portion when MUC1 is cleaved and the tandem repeat domain is shed. MNC2 and MNE6 will bind to a MUC1\*-like protein if it is devoid of tandem repeats, for example if a MUC1 negative cell is transfected or transduced with an engineered MUC1 that is devoid of tandem repeats, especially if extra cellular domain comprises the PSMGFR. Thus, the ectopic site to which MNC2 and MNE6 bind is unmasked when tandem repeat domain is missing or removed. Both MNC2 and MNE6 require the 10 membrane proximal amino acids of a MUC1\* extra cellular domain for binding; they do not bind to the C-10 peptide. That means that the ectopic binding site for MNC2 and MNE6 is within or contains all or part of the 10 C-terminal amino acids of the PSMGFR: PFPFSAQSGA. In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the inability of the antibody to bind to the C-10 peptide. In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to bind to the N-10 peptide and the inability of the antibody to bind to the C-10 peptide.

**[00690] Compete with NME7<sub>AB</sub> or NME7-X1 for binding to MUC1\* positive cell, PSMGFR peptide or N-10 peptide**

We have demonstrated that cancer-specific antibodies MNC2 and MNE6 bind to an ectopic epitope that comprises all or part of the 10 C-terminal amino acids of the PSMGFR peptide: PFPFSAQSGA. We have shown that growth factors, dimeric NME1 and NME7<sub>AB</sub>, also bind to an ectopic epitope that comprises all or part of the 10 C-terminal amino acids of the PSMGFR peptide. MNC2 and MNE6 compete with dimeric NME1 or NME7<sub>AB</sub> for binding to the PSMGFR peptide and the N-10 peptide. In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to disrupt the binding of NME1, NME7<sub>AB</sub>, or NME7-X1 to the PSMGFR peptide, the N-10 peptide, or to the surface of a MUC1\* positive cancer cell.

**[00691] Recognize a conformational epitope rather than a linear epitope**

**[00692]** Antibodies that are cancer-specific will be chosen based on their ability to bind to a MUC1 that is devoid of tandem repeats and for their inability to bind to full-length MUC1. Most often, MUC1\* is generated when MUC1 is cleaved by a cleavage enzyme and the tandem repeat domain is released from the cell surface. Cleavage and release of the tandem repeat domain may also unmask portions of MUC1\*-like cleavage products that exist on normal tissues. However, antibodies that recognize a conformation, rather than a linear epitope, are more selective. Antibodies that recognize a conformational epitope rather than a linear epitope can be identified by a variety of means. In particular, antibodies that recognize a conformational epitope will not work in a denaturing Western blot assay. In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to recognize a conformational epitope.

**[00693] Recognize a MUC1\* generated by cleavage by MMP9 or other tumor-associated cleavage enzyme**

**[00694]** The fold, or conformation, of the MUC1\* truncated extra cellular domain differs depending on which enzyme cleaves MUC1. Cleaved MUC1\* or MUC1\*-like cleavage products can function as growth factor receptors on normal healthy tissues. More than one cleavage enzyme is able to cleave MUC1 to a MUC1\*-like form. Cleavage by first enzyme may produce a conformation or a fold that is not the same as that produced by cleavage by a second enzyme. Support for this can be found in this application and is illustrated in Figures 39-41. These figures show that although a polyclonal antibody that binds to PSMGFR recognizes a cleaved MUC1 on hematopoietic stem cells, some monoclonal antibodies that bind to the PSMGFR peptide can bind to this MUC1\*-like form on hematopoietic stem cells while others cannot. For example, MNC3 readily recognizes this cleaved form of MUC1 on hematopoietic stem cells, but MNC2 and MNE6 do not. We know that MNC2 and MNE6 recognize a MUC1\* that is produced by cleavage by MMP9 but not when it is cleaved by MMP2. MNC2 and MNE6 are cancer-specific while MNC3 is not, as it recognizes stem and progenitor cells. We also know that MMP9 is overexpressed in cancers. Bone marrow, where hematopoietic stem cells are made expresses nearly 2,500-times more MMP2 than MMP9 (Fig. 65). MMP14 is another enzyme that cleaves MUC1 to a MUC1\* growth factor receptor form (Fig. 38). In one aspect of the invention, an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the

antibody to recognize a MUC1 cleavage product generated when MUC1 is cleaved by MMP14. In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to recognize a MUC1 cleavage product generated when MUC1 is cleaved by MMP9. In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to recognize a MUC1 cleavage product generated when MUC1 is cleaved by MMP9 and also recognizes a conformational epitope.

**[00695] Binds to cancer cells more than normal cells**

**[00696]** A traditional approach to identifying antibodies that are cancer-specific involves testing a panel of antibodies against a panel of different cancer cell lines and determining, by FACS, IF, immunoprecipitation or other method, if the antibody binds to cancer cells. Although this approach is traditional, it is sequential and time-consuming, and thus limits the analysis of large numbers of monoclonal antibody clones, which is required to find an ideal antibody suitable for cancer therapeutic or diagnostic. In addition, there are no real normal cell lines and the selection of normal primary cells is limited. The selection criteria presented above provide a rapid, multiplexed method for identifying monoclonal antibody clones that are specific for MUC1\* positive cancers. For many of the selection criteria, hybridoma supernatants can be used. This provides a huge advantage over state of the art methods for identifying antibodies that are specific for MUC1\* positive cancers. The ability to select antibodies from assay performed using the impure hybridoma supernatants means that much of the selection can be done on hundreds or thousands of clones rapidly and at very little cost. Methods such as FACS analysis and IHC tissue studies require the use of purified antibodies which limits the number of clones that can be tested to tens, not even hundreds.

**[00697]** However, selecting an antibody based on its ability to bind to cancer cells, or a cancer cell type or to a cell engineered to express a certain antigen is important for antibody selection. In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to bind to MUC1\* positive cancer cells.

**[00698] Binds to tumor tissue more than normal tissue**

**[00699]** Immunohistochemistry, IHC, tissue studies of cancerous versus normal tissues is a more stringent test of the cancer specificity of antibodies than FACS analysis. Cancer cell lines are a single cell from a single patient that have been expanded in a lab for decades and are not

representative of a cross section of the human population. Further, analysis of cell lines is blind to the heterogeneity of actual tumors. Tissue studies require purified antibody, are very expensive, time-consuming and require a skilled pathologist to analyze each stained tissue specimen. However, antibody staining of tissues from normal tissues versus cancerous tissues can reveal which antibodies cannot be used as therapeutics or diagnostics because of their cross-reactivity with normal tissues. Our systematic studies of numerous antibodies with thousands of human normal tissues or cancerous tissues, across several cancer sub-types showed that antibodies that bind to N-10, not C-10, disrupt the binding of NME1 or NME7<sub>AB</sub>, or NME7-X1 to the PSMGFR peptide, the N-10 peptide, or to the surface of a MUC1\* positive cancer cell, recognize a conformational epitope, and recognize a conformational epitope created by cleavage by MMP9 are the most cancer-specific.

**[00700]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to bind to MUC1\* positive tumor tissue at least 2-times more than it binds to normal tissues. In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to bind to MUC1\* positive tumor tissue at least 5-times more than it binds to normal tissues. In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on the ability of the antibody to bind to MUC1\* positive tumor tissue at least 10-times more than it binds to normal tissues.

**[00701] Antibodies that bind to refined epitopes**

**[00702]** In a preferred embodiment, an antibody, or fragments thereof, that binds to a peptide comprising the sequence QFNQYKTEAASRYNLTISDVSVSDVPFPFSAQSGA are incorporated into anti-cancer therapeutics or diagnostics.

**[00703]** In a more preferred embodiment, an antibody, or fragments thereof, that binds to a peptide comprising the sequence ASRYNLTISDVSVSDVPFPFSAQSGA are incorporated into anti-cancer therapeutics or diagnostics.

**[00704]** In a yet more preferred embodiment, an antibody, or fragments thereof, that binds to a peptide comprising the sequence SDVSVSDVPFPFSAQSGA are incorporated into anti-cancer therapeutics or diagnostics.

[00705] In a still more preferred embodiment, an antibody, or fragments thereof, that binds to a peptide comprising the sequence SVSDV are incorporated into anti-cancer therapeutics or diagnostics.


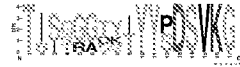



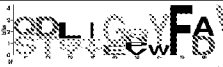
[00706] In a yet still more preferred embodiment, an antibody, or fragments thereof, that binds to a peptide comprising some or all of the sequence PFPFSAQSGA are incorporated into anti-cancer therapeutics or diagnostics.

#### [00707] Consensus Sequences

[00708] Antibodies of the invention were categorized according to cognate epitope. Sequences of their respective heavy chain CDRs are shown in **Table 5**. Sequences of their respective light chain CDRs are shown in **Table 6**. Consensus sequences for CDR1, CDR2 and CDR3 for each epitope-specific set of antibodies were computer generated. Figure 215 and Figure 216 show how the CDR consensus sequences change as the position of the antibodies' cognate epitope moves from the membrane-proximal portion of PSMGFR toward the more distal portions.

[00709] As can be seen in Table 5 and Table 6, the sequences for CDR1 and CDR2 for antibodies that bind to epitopes within the 10 membrane-proximal (C-terminal) portion of PSMGFR peptide closely adhere to the consensus sequence.

[00710] Table 5: HEAVY CHAIN CDRs

GTINVHVDVETQFNQYKTEAASRYNLTISDVSVDVPPFSAQSGA				
Epitope	Name	CDR1	CDR2	CDR3
<b>FPFS</b>	MNC2	FTFSGYAMS	TISSGGTYIYYPDSVKG	-LGGDNYEYFDV--
FPFS	MNE6	FTFSRYGMS	TISGGGTIYIYPDSVKG	DNYGRNYDYGMDY--
FPFS	20A1 0	FTFSTYAMS	-SIGRAGSTIYSDSVKG	---GPIYNDYDEFAY
FPFS	3C2B 1	ITFSTYTMS	TISTGGDKTYISDSVKG	-GTTAMYYYAMDY-
<b>Consensus Sequence</b>				
<b>SVSDV</b>	5C6F 3	FTFSTYAMS	AISNGGGYTYYPDSLKG	RYYDHYFDY
<b>ASRYNLT</b>	25E6	FTFSSYGMS	TISNGGRHTFYPSVKG	QTGTEGWFA Y
ASRYNLT	MNC3	YRFTDYAMN	VISTFSGNTNFNQKFKG	SDYYGPYFDY
ASRYNLT	18G1 2	YTFTGYFLY	GINPDNGGIDFNEKFRN	--LIGNY---
ASRYNLT	28F9	YTFTGYFLY	GIHPSNGD TDFNEKFKN	--LIGVY---
<b>Consensus Sequence</b>				
<b>QFNQYKTEA</b>	1E4	YAFSTYWMN	QIYPGSDTNYNGKFKG	GNHASMDY

GTINVHDTVET	18B4	FTFNDAWMD	EIRSTANIHTTYAESVQ G	-----LLYGFAY
GTINVHDTVET	29H1	FTFSDAWMD	EIRSKATNHATYYAESVK G	-----LLYGFAY
GTINVHDTVET	31A1	YTFTSYWMH	-- YINPSTGYTEYNQKFKD	-----AYIDY--
GTINVHDTVET	32C1	FTFSNYWMN	EIRLKSNNYAIHYAESVK G	VPGLDAY-----
GTINVHDTVET	39H5	YTFTNYGMN	-- WINTYTGEPTYVGDFKG	--GIHGYVDY--
GTINVHDTVET	3C5	YTFTNYGMN	-- WINTYTGKPTYADDFKG	-GGLDGYGYG-
Consensus Sequence				

Table 6: LIGHT CHAIN CDRs

GTINVHDTVETQFNQYKTEAASRYNLTISDVSVSDVPFPFSAQSGA				
Epitope	Name	CDR1	CDR2	CDR3
FPFS	MNC2	RASKS--VSTSGYSYMH	LASNLES	QHSRELPFT
FPFS	MNE6	-----SATSSVSYIH	STSNLAS	QQRSSSPFT
FPFS	20A10	KSSQSVLYSSNQKNYLA	WASTRES	-HQYLSLST
FPFS	3C2B1	RASKS--- ISTSDYNYIH	LASNLES	QHSRELPLT
Consensus Sequence				
SVSDV	5C6F3	RSSQTIVHSNGNTYLE	KVSNRFS	FQDSHVPLT
ASRYNLT	25E6	KSSQSLLDSDGKTYLN	LVSKLDS	WQGTTFPQT
ASRYNLT	MNC3	RSSQTIVHSNGNTYLE	KVSNRFS	FQGSHPVPT
ASRYNLT	18G12	KSSQSLHSDGKTYLI	LVSKLDS	CQGTTFPWT
ASRYNLT	28F9	KSSQSLHSDGKTYLI	LVSKLDS	CQGTTFPWT
Consensus Sequence				
QFNQYKTEA	1E4	RSSQSLVHSNGNTYLH	KVSNRFS	SQKTHVPWT
GTINVHDTVET	18B4	RTSQSLVHSNGNTYLH	KVSSRFS	SQNTTHVPYT
GTINVHDTVET	29H1	RSGQSLVHSNGHTYLH	KVSNRFS	SQTHVPWT
GTINVHDTVET	31A1	RSSQSIIVHSNGNTYLE	KVSNRFS	FQVSHFPWT
GTINVHDTVET	32C1	RSSQSLVHSNGNTYLH	KVSNRFS	SQITHVPYT
GTINVHDTVET	39H5	RSSQSIIVHRNGNTYL-	KVSNRFS	FQGSHPWT
GTINVHDTVET	3C5	KSSQSLHSGKTYLN	LVSKLES	LQTHFPWT
Consensus Sequence				

[00711] Whereas Heavy Chain CDR1 for MNC2 is FTFSGYAMS, with the amino acids numbered from left to right 1 through 9, the consensus of other antibodies that bind to that portion of PSMGFR is: F or I at position 1, T at position 2, F at position 3, S at position 4, T, G, or R at position 5, Y at position 6, A, G or T at position 7, M at position 8 and S at position 9.



**[00712]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a heavy chain CDR1 that is at least 90% identical to a CDR1 comprising the following amino acids at the specified positions: F or I at position 1, T at position 2, F at position 3, S at position 4, T, G, or R at position 5, Y at position 6, A, G or T at position 7, M at position 8 and S at position 9.

**[00713]** Whereas Heavy Chain CDR2 for MNC2 is TISSGGTYIYYPDSVKG, with the amino acids numbered from left to right 1 through 17, the consensus of other antibodies that bind to that portion of PSMGFR is: T at position 1, I or S at position 2, I or S at position 3, G or R at position 5, G or A at position 6, T or I at position 9, Y at position 10, Y at position 11, P or S at position 12 and DSVKG for positions 13-17.

**[00714]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a heavy chain CDR2 that is at least 90% identical to a CDR2 comprising the following amino acids at the specified positions: T at position 1, I or S at position 2, I or S at position 3, G or R at position 5, G or A at position 6, T or I at position 9, Y at position 10, Y at position 11, P or S at position 12 and DSVKG for positions 13-17.

**[00715]** Whereas Heavy Chain CDR3 for MNC2 is -LGGDNYEYFDV--, with the amino acids numbered from left to right 1 through 15, the consensus of other antibodies that bind to that portion of PSMGFR is: G, L, or N at position 2, G or T at position 4, Y at position 7, D or E at position 12, A at position 14, and Y at position 15.

**[00716]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a heavy chain CDR3 that is at least 90% identical to a CDR3 comprising the following amino acids at the specified positions: G, L, or N at position 2, G or T at position 4, Y at position 7, D or E at position 12, A at position 14, and Y at position 15.

**[00717]** Whereas Light Chain CDR1 for MNC2 is RASKS--VSTSGYSYMH, with the amino acids numbered from left to right 1 through 17, the consensus of other antibodies that bind to that portion of PSMGFR is: K or R at position 1, A or S at position 2, S at position 3, K or Q at position 4, S at position 5, V at position 6, L at position 7, T or S at position 10, Y at position 15, and I, L or M at position 16.

**[00718]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a light chain CDR1 that is at least 90% identical to a CDR1 comprising the following amino acids at the specified positions: K or R at position 1, A or S at

position 2, S at position 3, K or Q at position 4, S at position 5, L or V at position 6, L at position 7, T or S at position 10, Y at position 15, and I, L or M at position 16.

**[00719]** Whereas Light Chain CDR2 for MNC2 is LASNLES, with the amino acids numbered from left to right 1 through 7, the consensus of other antibodies that bind to that portion of PSMGFR is: L or W, or S at position 1, A or T at position 2, S at position 3, N or T at position 4, L or R at position 5, E or A at position 6, and S at position 7.

**[00720]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a light chain CDR2 that is at least 90% identical to a CDR2 comprising the following amino acids at the specified positions: L or W, or S at position 1, A or T at position 2, S at position 3, N or T at position 4, L or R at position 5, E or A at position 6, and S at position 7.

**[00721]** Whereas Light Chain CDR3 for MNC2 is QHSRELPFT, with the amino acids numbered from left to right 1 through 9, the consensus of other antibodies that bind to that portion of PSMGFR is: Q at position 1, H or Q at position 2, S, Q or R at position 3, R, S or Y at position 4, E, L, or S at position 5, L or S at position 6, P or S at position 7, F or L at position 8 and T at position 9.





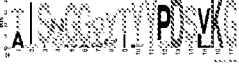

**[00722]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a light chain CDR3 that is at least 90% identical to a CDR3 comprising the following amino acids at the specified positions: Q at position 1, H or Q at position 2, S, Q or R at position 3, R, S or Y at position 4, E, L, or S at position 5, L or S at position 6, P or S at position 7, F or L at position 8 and T at position 9.

**[00723]** Another set of antibodies was generated and resultant clones were tested for their ability to bind to PSMGFR, N-10 and C-10 peptides. Antibody clones that bound to PSMGFR and N-10 peptides, but not to the C-10 peptide were selected. These antibodies were sequenced. Table 7 shows the sequences of the heavy chain CDRs for cancer-specific antibodies MNC2, MNE6, 20A10, 3C2B1, plus new antibodies B2, B7, 8C7F3, H11 and B9. Table 8 shows the sequences of the light chain CDRs for cancer-specific antibodies MNC2, MNE6, 20A10, 3C2B1, plus new antibodies B2, B7, 8C7F3, H11 and B9. Consensus sequences for the heavy and light chain CRDs were generated and are shown in Table 7 and Table 8. Although antibodies 5C6F3 and 25E6 showed great cancer specificity in IHC tissue studies and they both bound to the PSMGFR and N-10 peptides, but not to the C-10 peptide, epitope mapping showed that they

bound to epitopes that were a bit N-terminal to the epitopes to which MNC2, MNE6, 20A10 and 3C2B1 bound. For this reason, consensus sequences were generated for MNC2, MNE6, 20A10, 3C2B1 and the new antibodies plus consensus sequences were generated for all the antibodies that bound to N-10 but not to C-10.






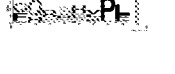
**[00724]** As can be seen in Table 7 and Table 8, the sequences for CDR1, CDR2 and CDR3 for antibodies that require for binding the 10 membrane-proximal (C-terminal) amino acids of PSMGFR peptide closely adhere to a common consensus sequence.

**[00725] Table 7: HEAVY CHAIN CDRs for antibodies that share broader epitope in that they cannot bind to the C-10 peptide**

GTINVHVDVETQFNQYKTEAASRYNLTISDVSVDVFPFSAQSGA				
Epitope	Name	CDR1	CDR2	CDR3
FPFS	MNC2	FTFSGYAMS	TISSGGTYIYPDS VKG	-LGGDNYEYFDV--
FPFS	MNE6	FTFSRYGMS	TISGGGTYYYPDS VKG	DNYGRNYDYGMDY--
FPFS	20A10	FTFSTYAMS	- SIGRAGSTYYSDSV KG	---GPIYNDYDEFAY
FPFS	3C2B1	ITFSTYTMS	TISTGGDKTYSDS VKG	-GTTAMYYYAMDY--
PFPSAQSGA	B2	FAFSTFAMS	AISNGGGYTYYPDT LKG	----RYYDLYFDL--
PFPSAQSGA	B7	FTFSRYGMS	TISSGGTYIYPDS VKG	DNYGSSYDYAMDY--
PFPSAQSGA	8C7F3	FTFSTYAMS	AISNGGGYTYYPDS LKG	----RYYDHYFDY--
PFPSAQSGA	H11	FAFSTFAMS	AISNGGGYTYYPDT LKG	----RYYDLYFDL--
PFPSAQSGA	B9	FTFSRYGMS	TISSGGTYIYPDS VKG	DNYGSSYDYAMDY--
Consensus Sequence - all of antibodies above				
SVSDV	5C6F3	FTFSTYAMS	AISNGGGYTYYPDS LKG	RYYDHYFDY
ASRYNLT	25E6	FTFSRYGMS	TISNGGRHTFYPS VKG	QTGTEGWFAFAY
Consensus Sequence - all antibodies				

**Table 8: LIGHT CHAIN CDRs for antibodies that share broader epitope in that they cannot bind to the C-10 peptide**

GTINVHVDVETQFNQYKTEAASRYNLTISDVSVDVFPFSAQSGA				
Epitope	Name	CDR1	CDR2	CDR3
FPFS	MNC2	RASKS-- VSTSGYSYMH	LASNLES	QHSRELPFT
FPFS	MNE6	----- SATSSVSYIH	STSNLAS	QQRSSSPFT

FPFS	20A10	KSSQSVLYSSNQKNYL A	WASTRES	-HQYLSSLT
FPFS	3C2B1	RASKS--- ISTSDYNYIH	LASNLES	QHSRELPLT
<b>PFPSAQSGA</b>	B2	RSSQNIV- HSNGNTYLE	KVSNRFS	FQDSHVPLT
<b>PFPSAQSGA</b>	B7	RSSQTIV- HSNGNTYLE	KVSNRFS	FQDSHVPLT
<b>PFPSAQSGA</b>	8C7F3	-- RASESVATYGNNFMQ	LASTLDS	QQNNEDPPT
<b>PFPSAQSGA</b>	H11	RSSQNIV- HSNGNTYLE	KVSNRFS	FQDSHVPLT
<b>PFPSAQSGA</b>	B9	----- SASSSVSYM	TTSNLAS	QQRSSYPF-
<b>Consensus Sequence - all of antibodies above</b>				
<b>SVSDV</b>	5C6F3	RSSQTIVHSNGNTYLE	KVSNRFS	FQDSHVPLT
<b>ASRYNLT</b>	25E6	KSSQSLLDSDGKTYLN	LVSKLDS	WQGFHFPQT
<b>Consensus Sequence - all antibodies</b>				

[00726] Whereas Heavy Chain CDR1 for MNC2 is FTFSGYAMS, with the amino acids numbered from left to right 1 through 9, the consensus sequence of MNC2, MNE6, 20A10, 3C2B1 and new antibodies B2, B7, 8C7F3, H11 and B9 is: F or I at position 1, T or A at position 2, F at position 3, S at position 4, T, G, or R at position 5, Y or F at position 6, A, G or T at position 7, M at position 8 and S at position 9. The underlined amino acids at positions 2 and 6 are the only additional variants to the consensus sequence generated for cancer-specific antibodies MNC2, MNE6, 20A10, 3C2B1 alone.

[00727] As can be seen in Table 7, the inclusion of antibodies 5C6F3 and 25E6 into the generation of consensus sequence did not change in any way the consensus sequence for heavy chain CDR1 that describes a cancer-specific anti-MUC1\* antibody.

[00728] In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a heavy chain CDR1 that is at least 90% identical to a CDR1 comprising the following amino acids at the specified positions: F or I at position 1, T or A at position 2, F at position 3, S at position 4, T, G, or R at position 5, Y or F at position 6, A, G or T at position 7, M at position 8 and S at position 9.

**[00729]** Whereas Heavy Chain CDR2 for MNC2 is TISSGGTYIYYPDSVKG, with the amino acids numbered from left to right 1 through 17, the consensus sequence of MNC2, MNE6, 20A10, 3C2B1 and new antibodies B2, B7, 8C7F3, H11 and B9 is:

**[00730]** T or A at position 1, I or S at position 2, I or S at position 3, N, S, T or G at position 4, G or R at position 5, G or A at position 6, G, T, or D at position 7, Y, K or S at position 8, T or I at position 9, Y at position 10, Y at position 11, P or S at position 12 and D at position 13, S or T at position 14, V or L at position 15 and KG for positions 16-17. The underlined amino acids indicate how this more inclusive consensus sequence differs from the consensus sequence generated for MNC2, MNE6, 20A10 and 3C2B1 alone. Of the 17 amino acids in heavy chain CDR2, the consensus sequence for all nine antibodies differs from the consensus sequence for the original cancer-specific four by only 4 amino acids. Note that 2 of the 4 variants are homologous changes, T for S and L for V, which generally do not significantly impact the structure or specificity of a protein.

**[00731]** As can be seen in Table 7, the inclusion of antibodies 5C6F3 and 25E6 into the generation of consensus sequence for heavy chain CDR2 only changed the consensus sequence by the addition of two other possible amino acids: a possible H at position 8, and a possible F at position 10, for a heavy chain CDR2 that describes a cancer-specific anti-MUC1\* antibody. We note that the change of Y to F at position 10 is a homologous change, which generally does not significantly impact the structure or specificity of a protein.

**[00732]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a heavy chain CDR2 that is at least 90% identical to a CDR2 comprising the following amino acids at the specified positions: T or A at position 1, I or S at position 2, I or S at position 3, N, S, T or G at position 4, G or R at position 5, G or A at position 6, G, T, or D at position 7, Y, K, H or S at position 8, T or I at position 9, Y or F at position 10, Y at position 11, P or S at position 12 and D at position 13, S or T at position 14, V or L at position 15 and KG for positions 16-17.

**[00733]** Whereas Heavy Chain CDR3 for MNC2 is LGGDNYEYFDV, with the amino acids numbered from left to right 2 through 13, the consensus sequence of MNC2, MNE6, 20A10, 3C2B1 and new antibodies B2, B7, 8C7F3, H11 and B9 is:

**[00734]** G, L, or N at position 2, G, T, or Y at position 3, G or T at position 4, A, D, P, R, or S at position 5, Y, M, I or S at position 6, Y at position 7, D, Y, or N at position 8, E, D, Y, L or

H at position 9, Y, A, or G at position 10, M, D or F at position 11, D or E at position 12, V, F, Y or L at position 13, and AY at position 14-15. The underlined amino acids indicate how this more inclusive consensus sequence differs from the consensus sequence generated for MNC2, MNE6, 20A10 and 3C2B1 alone. Of the 15 amino acids in heavy chain CDR3, the consensus sequence for all nine antibodies differs from the consensus sequence for the original cancer-specific four by 7 amino acids, with 3 of the 7 substitutions at position 6. For this reason, we conclude that the amino acid at position 6 can be varied without altering the specificity of the antibody.

**[00735]** Analysis of the consensus sequence generated with the inclusion of antibodies 5C6F3 and 25E6 highlighted which amino acids were conserved among all eleven antibodies. For this reason, our preferred consensus sequence for heavy chain CDR3 defines amino acids at positions 2, 3, 4, 7, 10, 11, 12, 14 and 15, where for 11 antibodies, there were 3 or less variants at these positions.

**[00736]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a heavy chain CDR3 that is at least 90% identical to a CDR3 comprising the following amino acids at the specified positions: G, L, or N at position 2, G, T, or Y at position 3, G or T at position 4, Y at position 7, Y, A, or G at position 10, M, D or F at position 11, D or E at position 12 and AY at position 14-15.

**[00737]** Whereas Light Chain CDR1 for MNC2 is RASKS--VSTSGYSYMH, with the amino acids numbered from left to right 1 through 17, the consensus sequence of MNC2, MNE6, 20A10, 3C2B1 and new antibodies B2, B7, 8C7F3, H11 and B9 is:

**[00738]** K or R at position 1, A or S at position 2, S or R at position 3, K, Q or A at position 4, S, N or T at position 5, V, I, E, or K at position 6, L, V or S at position 7, S, Y, I or V at position 8, A, S, or H at position 9, T or S at position 10, N, S, or Y at position 11, G, S, D, or Q at position 12, V, Y, K or N at position 13, N, S, or T at position 14, Y or F at position 15, and I, L or M at position 16, and H, A, E or Q at position 17. The underlined amino acids indicate how this more inclusive consensus sequence differs from the consensus sequence generated for MNC2, MNE6, 20A10 and 3C2B1 alone. Of the 17 amino acids in light chain CDR1, the consensus sequence for all nine antibodies differs from the consensus sequence for the original cancer-specific four by 13 amino acids. 4 of the 13 are homologous substitutions, which in general do not significantly alter the structure or specificity of the protein. Of the remaining 9

substitutions, 1 is at position 4, 1 is at position 5, 3 are at position 6, 1 is at position 7, 1 is at position 11, and 2 are at is at position 17. The inclusion of the 5 new antibodies did not alter the amino acids, excluding homologous substitutions, at positions 1, 2, 3, 8, 9, 10, 12, 13, 14, 15 or 16. For this reason, we conclude that the conserved consensus sequence for light chain CDR1 that defines a MUC1\* cancer-specific antibody comprises the amino acids given above for positions 1, 2, 3, 8, 10, 12, 13, 14, 15 and 16.

**[00739]** Analysis of the consensus sequence generated with all the antibodies, including 5C6F3 and 25E6 further altered the consensus sequence for light chain CDR1 with amino acid substitutions as follows: L at position 6; D at position 9; D at position 11 and N at position 17. We note that none of these substitutions were at positions that were invariant for the original four cancer-specific antibodies plus the five new antibodies. Thus, we conclude that a conserved consensus sequence for light chain CDR1 that defines at least 90% identity of a cancer-specific antibody comprises amino acids defined above at positions 1, 2, 3, 8, 10, 12, 13, 14, 15 and 16.

**[00740]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a light chain CDR1 that is at least 90% identical to a CDR1 comprising K or R at position 1, A or S at position 2, S or R at position 3, S, Y, I or V at position 8, T or S at position 10, G, S, D, or Q at position 12, V, Y, K or N at position 13, N, S, or T at position 14, Y or F at position 15, and I, L or M at position 16.

**[00741]** Whereas Light Chain CDR2 for MNC2 is LASNLES, with the amino acids numbered from left to right 1 through 7, the consensus sequence of MNC2, MNE6, 20A10, 3C2B1 and new antibodies B2, B7, 8C7F3, H11 and B9 is: L, W, S, T or K at position 1, A, T or V at position 2, S at position 3, N or T at position 4, L or R at position 5, E, A, F or D at position 6, and S at position 7. The underlined amino acids indicate how this more inclusive consensus sequence differs from the consensus sequence generated for MNC2, MNE6, 20A10 and 3C2B1 alone.

**[00742]** In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a light chain CDR2 that is at least 90% identical to a CDR2 comprising the following amino acids at the specified positions: L, W, S, T or K at position 1, A, T or V at position 2, S at position 3, N or T at position 4, L or R at position 5, E, A, F or D at position 6, and S at position 7. Of the 7 positions, the inclusion of the five new antibodies introduced 5 substitutions of which only 2 were not homologous substitutions.



[00743] Analysis of the consensus sequence generated with all the antibodies, including 5C6F3 and 25E6 further altered the consensus sequence for light chain CDR2 with amino acid substitutions as follows: K at position 4, which is a substitution that is homologous to N.

[00744] In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a light chain CDR2 that is at least 90% identical to a CDR2 comprising: A, T or V at position 2, S at position 3, N, T, or K at position 4, L or R at position 5, E, A, F or D at position 6, and S at position 7.

[00745] Whereas Light Chain CDR3 for MNC2 is QHSRELPFT, with the amino acids numbered from left to right 1 through 9, the consensus sequence of MNC2, MNE6, 20A10, 3C2B1 and new antibodies B2, B7, 8C7F3, H11 and B9 is: Q or F at position 1, H or Q at position 2, S, Q, R, D or N at position 3, R, S, Y or N at position 4, E, L, S or H at position 5, L, S, V, D or Y at position 6, P or S at position 7, F, L or P at position 8 and T at position 9. The underlined amino acids indicate how this more inclusive consensus sequence differs from the consensus sequence generated for MNC2, MNE6, 20A10 and 3C2B1 alone.

[00746] Analysis of the consensus sequence generated with all the antibodies, including 5C6F3 and 25E6 further altered the consensus sequence for light chain CDR2 with amino acid substitutions as follows: W at position 1; G at position 3; T at position 4; F at position 5; Q at position 8.

[00747] In a preferred embodiment an antibody is chosen for the treatment, prevention or diagnosis of cancer based on having a light chain CDR3 that is at least 90% identical to a CDR2 comprising: Q, F or W at position 1, H or Q at position 2, R, S, T, Y or N at position 4, E, L, S or H at position 5, L, S, V, D or Y at position 6, P or S at position 7, and T at position 9.

**[00748] Other general strategy for using antibodies, antibody fragments and CARs that target the extracellular domain of MUC1\***

[00749] In another aspect, the invention is directed to a composition that includes at least two different plasmids transfected into the same immune cell, wherein the first encodes a CAR comprising an antibody fragment, scFv, or peptide that binds to a tumor antigen and the other encodes a gene that is not a CAR, wherein the gene that is not a CAR is expressed from an inducible promoter that is activated by elements of an activated immune cell. In one aspect, the immune cell is a T cell or an NK cell. In one aspect the CAR comprises an antibody fragment, scFv or peptide that binds to the extra cellular domain of MUC1\*. In one aspect the CAR

comprises an scFv derived from MNC2, MNE6, 20A10, 3C2B1, 5C6F3, 25E6, 18G12, 28F9, 1E4, B12, B2, B7, B9, 8C7F3, or H11. In one aspect the non-CAR species is a cleavage enzyme. In one aspect the cleavage enzyme is MMP2, MMP3, MMP9, MMP13, MMP14, MMP16, ADAM10, ADAM17, ADAM28 or catalytically active fragments thereof. In another aspect the non-CAR species is a cytokine. In one aspect, the Cytokine is IL-7. In one aspect the cytokine is IL-15. In one aspect the cytokine is IL-12. In one aspect the cytokine is IL-18. The sequence of an activated IL-18 is given (SEQ ID NOS:1637-1638). Two examples of NFAT-inducible IL-18 embedded in the Foxp3 enhancer region are given (SEQ ID NOS:1639-1640). Two examples of NFAT-inducible IL-18 embedded in the IL-2 enhancer region are given (SEQ ID NOS:1641-1642). In one case, there are three (3) NFAT response elements and in the other case there are six (6) NFAT response elements. The number of NFAT response elements can be varied in order to get the desired amount of IL-18 expressed upon CAR T cell recognition of the target. Examples of antibodies of the invention incorporated into CARS with inducible IL-18 are shown as: murine or human MNC2 in a CAR with a 4-1BB or CD28 co-stimulatory domain plus inducible IL-18 (SEQ ID NOS:1643-1646), or also with a 1XX mutated CD3-zeta (SEQ ID NOS:1647-1650); murine or human MNE6 in a CAR with a 4-1BB or CD28 co-stimulatory domain plus inducible IL-18 (SEQ ID NOS:1651-1654), or also with a 1XX mutated CD3-zeta (SEQ ID NOS:1655-1658); murine or human 20A10 in a CAR with a 4-1BB or CD28 co-stimulatory domain plus inducible IL-18 (SEQ ID NOS:1659-1662), or also with a 1XX mutated CD3-zeta (SEQ ID NOS:1663-1666); murine or human 25E6 in a CAR with a 4-1BB or CD28 co-stimulatory domain plus inducible IL-18 (SEQ ID NOS:1667-1670), or also with a 1XX mutated CD3-zeta (SEQ ID NOS:1671-1674). In another aspect the cytokine is IL-7 and IL-15. In one case expression of the non-CAR species is induced by elements of an activated immune cell. In one aspect the element of an activated immune cell is an NFAT. In one aspect the NFAT is NFATc1, NFATc3 or NFATc2. Cytokines IL-7, IL-15, IL-12 and IL-18 are known to promote T cell persistence. In one aspect of the invention an immune cell described above is administered to a patient for the treatment or prevention of cancer. In one aspect of the invention, the cancer is a MUC1 positive cancer or a MUC1\* positive cancer.

**[00750]** In addition to making CAR T cells that also induce expression of a cleavage enzyme, we made CAR T cells that also induce local and transient expression of IL-18. Many of the T cell based inducible systems reported insert the gene to be inducibly expressed into an IL-2

promoter or enhancer. We compared inducible expression off an IL-2 promoter/enhancer to inducible expression off of a portion of the Foxp3 enhancer. In this particular example, human T cells were transduced with both huMNC2-CAR44 and an NFAT inducible IL-18, wherein the IL-18 gene was either inserted into an IL-2 promoter or the Foxp3 enhancer region. It is known in the field that a major problem with CAR Ts with inducible second factors is that the second factor is leaky, meaning that significant expression of the second factor occurs without activation of the CAR T cell. The other problem with existing inducible systems is the length of time that goes by between when the CAR T cell is activated and the second factor is induced is typically very long so that the cell secreting the second factor may be far away from the tumor by the time the second factor is expressed.

**[00751]** **Figure 211A- 211C** show graphs of an ELISA experiment measuring the amount of IL-18 secreted into the condition media of huMNC2-CAR44 T cells, which also bear an NFAT inducible IL-18, co-cultured with MUC1\* positive cancer cells. As a method of inducing varying levels of IL-18 expression, we co-cultured the CAR T cells with cancer cells doped with increasing amounts of cells that were engineered to express even more MUC1\*. In these figures we show T47D cancer cells that are either wild-type, or doped with 5%, 10% or 30% of the T47D cells expressing more MUC1\*. Fig. 211A shows the graph of IL-18 secreted into the supernatant of T47D breast cancer cells co-cultured with untransduced human T cells. Fig. 211B shows the graph of IL-18 secreted into the supernatant of T47D breast cancer cells co-cultured with huMNC2-CAR44 T cells that also bore an NFAT inducible IL-18 gene inserted into a portion of the Foxp3 enhancer. Fig. 211C shows the graph of IL-18 secreted into the supernatant of T47D breast cancer cells co-cultured with huMNC2-CAR44 T cells that also bore an NFAT inducible IL-18 gene inserted into a portion of the IL-2 enhancer. As can be seen in the figure, the Foxp3 system induces rapid and robust expression of IL-18, which is significantly faster and higher than that of the same construct in an IL-2 promoter. In this example, the IL-18 gene is inserted downstream of six (6) NFAT response elements, however one can attenuate the amount of the second factor by using a lesser number of response elements or enhance the amount by increasing the number of NFAT response elements.

**[00752]** It has been reported that IL-18 increases persistence of CAR T cells in vivo. However, we observed an unexpected result. In a dose-dependent manner, secretion of IL-18 increased the killing of low antigen density cells by the CAR T cells. We differentially labeled

the T47D-wt cells (red: mCherry) and those T47Ds that were transduced to express more MUC1\* (green: GFP). **Figure 212A- 212X** shows photographs of T47D breast cancer cells (red) doped with varying percentages of T47D cells engineered to express more MUC1\* (green). The target cancer cells have been co-cultured with huMNC2-CAR44 T cells with NFAT inducible IL-18 wherein the IL-18 gene has been inserted into either the Foxp3 enhancer/promoter or the IL-2 enhancer/promoter. Fig. 212A-212C, 212I-212K, and 212Q-212S show the cancer cells co-cultured with untransduced T cells. Fig. 212D-212F, 212L-212N, and 212T-212V show the cancer cells co-cultured with hiMNC2-CAR44 T cells with the NFAT inducible IL-18 gene inserted into the Foxp3 enhancer/promoter. Fig. 212G-212H, 212O-212P, and 212W-212X show the cancer cells co-cultured with hiMNC2-CAR44 T cells with the NFAT inducible IL-18 gene inserted into the IL-2 enhancer/promoter. As can be seen in the figure, the low antigen density T47D-wt type cells (red) are being killed when doped with higher percentages of cells that express more MUC1\* and thus secrete more IL-18. The experiment shows that this is not just a bystander effect, because the cells expressing IL-18 off of the IL-2 promoter, which expresses much lower levels of IL-18, do not kill the low antigen density cells even when they are doped with 30% cells expressing more MUC1\*.

**[00753]** We then showed that the CAR T mediated killing is specific for the CAR T specific antigen. We performed a similar experiment, wherein control, MUC1/MUC1\* negative cells were doped with 5%, 10% or 30% of the T47D cells expressing more MUC1\*, and co-cultured with MUC1\* specific CAR T cells. **Figure 213A- 213B** shows graphs of ELISA experiments in which levels of IL-18 secreted into the conditioned media are measured for huMNC1-CAR44 T cells with NFAT inducible IL-18 gene, inserted into the Foxp3 enhancer or promoter, co-cultured with either MUC1\* positive cancer cells or MUC1 negative non-cancerous cells. Fig. 213A shows IL-18 secretion from huMNC2-CAR44 T cells with NFAT inducible IL-18 in co-culture with T47D breast cancer cells where the population has been doped with 5%, 10% or 30% T47D cells that had been transfected with even more MUC1\*. Fig. **213B** shows IL-18 secretion from huMNC2-CAR44 T cells with NFAT inducible IL-18 in co-culture with non-cancerous, MUC1 negative HEK293 cells where the cell population has been doped with 5%, 10% or 30% T47D cells that had been transfected with more MUC1\*. As can be seen in the figure, the amount of IL-18 secreted into the media can be attributed to the MUC1\* positive cells that the population was doped with. Time course fluorescent photographs of the experiment

show that even when doped with significant percentages of high antigen density MUC1\* positive cells, the MUC1 negative cells are not killed by the MUC1\* targeting CAR T cells. **Figure 214A-214X** shows photographs of T47D breast cancer cells (red) or non-cancerous HEK293 cells (also red), where both cell types have been doped with varying percentages of T47D cells engineered to express more MUC1\* (green). These target cancer cells have been co-cultured with huMNC2-CAR44 T cells with NFAT inducible IL-18 wherein the IL-18 gene has been inserted into the Foxp3 enhancer/promoter. Fig. 214A-214F shows either T47D cells or HEK293 cells that have not been doped with T47D cells engineered to express high MUC1\* density. Fig. 214G-214L shows either T47D cells or HEK293 cells that have been doped with 5% T47D cells engineered to express high MUC1\* density. Fig. 214M-214R shows either T47D cells or HEK293 cells that have been doped with 10% T47D cells engineered to express high MUC1\* density. Fig. 214S-214X shows either T47D cells or HEK293 cells that have been doped with 30% T47D cells engineered to express high MUC1\* density. Fig. 214A-B, G-H, M-N, and S-T show T47D breast cancer cells. Fig. 214C-F, I-L, O-R, and U-X show HEK293 cells. As can be seen in the figures, the induced secretion of IL-18 resulted in low MUC1\* density T47D cells being killed but did not induce non-specific killing of the MUC1\* negative HEK293 cells. Taken together these results show that the Foxp3 system is a superior system for the inducible expression of a second factor and especially useful in CAR T systems. Further we have demonstrated the unexpected result that IL-18 increases the killing of low antigen density cells without the unwanted effect of killing nearby MUC1/MUC1\* negative cells.

**[00754]** In another aspect, the invention is directed to a composition that includes at least two different plasmids transfected into the same immune cell, wherein the first encodes a CAR comprising an antibody fragment, scFv or peptide that binds to the extra cellular domain of an antigen on the surface of a B cell and the other encodes a gene that is not a CAR, wherein the gene that is not a CAR is expressed from an inducible promoter that is activated by elements of an activated immune cell. In one aspect, the immune cell is a T cell or an NK cell. In one aspect the CAR comprises an antibody fragment, scFv or peptide that binds to CD19. In another aspect the antibody fragment, scFv or peptide binds to a surface antigen of a B cell or a B cell precursor, or binds to CD19, CD20, CD22, BCMA, CD30, CD138, CD123, CD33 or LeY antigen. In one aspect the non-CAR species is a cleavage enzyme. In another aspect the non-CAR species is a cytokine. In one aspect, the Cytokine is IL-7. In one aspect the cytokine is IL-

15. In another aspect the cytokine is IL-7 and IL-15. In one case expression of the non-CAR species is induced by elements of an activated immune cell. In one aspect the element of an activated immune cell is an NFAT. In one aspect the NFAT is NFATc1, NFATc3 or NFATc2. that is not a CAR, wherein the gene that is not a CAR is expressed from an inducible promoter wherein expression is induced by elements of an activated immune cell. In one aspect the immune cell transfected or transduced with the composition is administered to a patient for the treatment or prevention of cancer. In one case the cancer is a leukemia, lymphoma or blood cancer.

**[00755]** It is not intended for the invention to be limited by a specific method or technology for inserting the gene or plasmid comprising a sequence encoding a CAR or activated T cell inducible protein or peptide there encoded. For example, the gene encoding the CARs and activated T cell induced genes described herein can be virally transduced into an immune cell using viruses, which may or may not result in the CAR gene being integrated into the genome of the recipient cell. Virus delivery systems and viral vectors include but are not limited to retroviruses, including gamma-retroviruses, lentivirus, adenoviruses, adeno-associated viruses, baculoviruses, poxvirus, herpes simplex viruses, oncolytic viruses, HF10, T-Vec and the like. In addition to viral transduction, CARs and activated T cell induced genes described herein can be directly spliced into the genome of the recipient cell using methods such as CRISPR technology, CRISPR-Cas9 and -CPF1, TALEN, Sleeping Beauty transposon system, and SB 100X.

**[00756]** Bulky cell surface proteins such as MUC1-FL can also cause a steric hindrance problem for BiTEs. A BiTE is a two-headed bi-specific antibody wherein one head binds to a T cell and the other head binds to a tumor-associated antigen. In this way, the BiTE links together the T cell and the tumor cells. The antibody that binds to the T cell should be an antibody that activates the T cell, such as an antibody against CD3 or CD28. To solve the steric hindrance problem, the linker between the T cell specific antibody and the tumor specific antibody is lengthened.

**[00757]** In another aspect of the invention, an anti-MUC1\* single chain molecule is fused to a cleavage enzyme or a catalytically active fragment of a cleavage enzyme. In one aspect of the invention, the cleavage enzyme is MMP9 (SEQ ID NO:643). In another aspect of the invention, the enzyme is a catalytically active fragment of MMP9 (SEQ ID NO:645). In some cases, the antibody fragment of the CAR is chosen for its ability to recognize MUC1\* when cleaved by that

specific cleavage enzyme. In one embodiment, the cleavage enzyme is MMP9, MMP3, MMP14, MMP2, ADAM17, ADAM TS16, and/or ADAM28. In one embodiment, the antibody or antibody fragment binds to a peptide having the sequence of (PSMGFR) GTINVHDTVETQFNQYKTEAASRYNLTISDVSVDVPPFSAQSGA, PSMGFR N-10, QFNQYKTEAASRYNLTISDVSVDVPPFSAQSGA, or PSMGFR N+20 SNIKFRPGSVVVQLTLAFREGTINVHDTVETQFNQYKTEAASRYNLTISDVSVDVPPFSAQSGA. In another embodiment, cleavage enzymes MMP9 and MMP3 are transduced into a T cell that is also transduced with a CAR with an antibody fragment that is a fragment of MNC2.

**[00758]** In many cases it is desirable to have the cleavage enzyme expressed only after an immune cell recognizes the tumor-associated target on a solid tumor. In this way, the cleavage enzyme will not freely move throughout the body, cleaving MUC1, MUC16 or other proteins, wherein their cleavage could actually promote cancer. However, there are cancers that are physically accessible to direct application of chemotherapy agents, CAR T cells and other anti-cancer agents. For example, types of brain cancers, prostate cancer and ovarian cancers have all shown the benefit of direct application of anti-cancer agents into the local vicinity of the cancer. CAR T cells have been injected directly into the brain and/or cerebral spinal fluid of glioblastoma patients. Radiation has been directed to the prostate area for the treatment of prostate cancers, including those that have metastasized. Hot chemo therapy agents have been directly injected into the intraperitoneal cavity for the treatment of ovarian cancers. In these and other cases, where the cancers that are physically accessible to direct application of chemotherapy agents, a cleavage enzyme is administered in the presence or absence of another anti-cancer agent, which could be a CAR T cell, an immune cell engineered to recognize a tumor-associated antigen, a BiTE, an ADC, a biological or a standard chemotherapy agent. Although ovarian cancer can metastasize to anywhere in the body, it usually stays in the abdomen as it spreads to adjacent organs, such as the intestines, liver and stomach. This makes ovarian cancer an ideal test case for improving the effect of anti-cancer agents by administering a cleavage enzyme in combination with other anti-cancer agents, including a platinum-based drug such as carboplatin (Paraplatin) or cisplatin, and/or a taxane such as paclitaxel (Taxol) or docetaxel (Taxotere). Alkeran (Melphalan), Avastin (Bevacizumab), Carboplatin, Clafen (Cyclophosphamide), and Cytosan have all been approved for the treatment of ovarian

cancer. Other treatments that are being tested for the treatment of ovarian cancers include agents that target MUC1, MUC16 and as described herein, MUC1\*.

**[00759]** Other cleavage enzymes can be used in addition to or in place of MMP9. MMP14 for example, has been shown to efficiently cleave MUC1 to MUC1\* (Fig. 38). In one aspect of the invention, MMP14 is expressed in an immune cell that is also engineered to express a CAR. In one case the CAR is an anti-MUC1\* CAR. For example, it can be an MNC2-CAR44 transduced T cell. In another aspect of the invention, the MMP14 is directly administered to the patient either in the location of the tumor or by i.v.

**[00760]** In yet another aspect of the invention, the cancer is an ovarian cancer and either MMP9 or MMP14 is directly injected into the abdominal area along with an anti-cancer agent, which can be a chemotherapy agent, a biological, an anti-MUC1\* CAR T or an anti-MUC16 CAR T.

**[00761]** In addition to local administration of the cleavage enzyme, + iv administration alone or secreted from an immune cell, which may be a CAR T cell, which further may be expressed off of an inducible promoter is contemplated.

**[00762] Methods used in carrying out experimentation in relation to the present invention**

**[00763] 1. *Lentivirus production and viral transduction of immune cells***

**[00764]** HEK293 or HEK293T cells (ATCC) were used to produce lentivirus. The day prior transfection plates (6well plate) were coated with poly-D-lysine and cells seeded so that cell density reaches 90-95% at the time of transfection and cultures in a 5% CO<sub>2</sub> atmosphere. The next day cells were transfected with Lipofectamine 3000 (life technologies) and Opti-MEM® I Reduced Serum Medium according to the manufacturer instructions (0.75ug of lentiviral expression vector and 2.25ug of pPACKH1 packaging mix was used). After 6h incubation, the media was changed and media containing lentivirus was harvested after 24 and 48 hours. Lentivirus was concentrated with Lenti-X concentrator (Clontech) and titer was calculated using the Lenti-X p@4 Rapid Titer Kit (Clontech). Lentivirus was store at -80C in single-use aliquots.

**[00765] Transduction of immune cells with constructs including CARs**

**[00766]** Human T cells, if frozen, were thawed and pre-warmed in 100-200 units IL-2 and TexMACS medium, 20 ml, and pelleted by centrifugation. Cells were resuspended in 10 ml of



medium and cultured at 37°C, 5% CO<sub>2</sub> at 1x10<sup>6</sup> cells/ml in complete medium with anti-CD3/anti-CD28 beads (TransAct kit).

**[00767]** After 4 days in culture, cells were counted and 450 ul of cell suspension was placed in single well of a 24-well plate at a density of approximately 1x10<sup>6</sup> cells/ml. Cells were allowed to settle. 150 ul was carefully removed from the top of each well. To each well was added an appropriate dilution of lentiviral vector, diluted in plain TexMACS medium, along with protamine sulfate to a final concentration of 10 ug/ml, in a 150 ul volume, for a final total volume of 450 ul per well and incubated for 24 hrs. Transduced cells were removed, pelleted by centrifugation, and resuspended in fresh medium, adjusting cell density, not to exceed 1.0 x 10<sup>6</sup> cells/ml. Transduced T cells can be expanded and frozen or used directly. Typically transduced T cells are used or frozen between Day 7 and Day 20 post activation with IL-2 and TransAct media.

**[00768] 2. Comparing anti-MUC1\* CAR T cell activity in the presence or absence of exogenous cleavage enzymes**

**[00769]** Human T cells (ALLCELLS) were transduced with huMNC2-CAR44 or huMNC2-CAR50. CAR44 is huMNC2-scFv-CD8-CD8 (transmembrane-41BB-3z). CAR50 is the same as CAR44 except that CAR50 has a murine MNC2-scFv and a CD4 transmembrane domain. The CAR T cells were incubated for 18 hours with target and non-target cells that have been dyed red using CMTMR. When T cells recognize a target cell, they cluster the target cells and begin to kill them. As can be seen in **Figures 45-47** the CAR T cells effectively cluster and kill the target MUC1\* positive cancer cells. Figure 45 shows huMNC2-CAR44 or huMNC2-CAR50 T cells being co-cultured with HCT-116 cells transduced to express MUC1\*, “HCT-MUC1\*” or with HCT-116 cells transduced with a full-length MUC1, “HCT-MUC1-41TR”. Recall that MNC2 recognizes an ectopic epitope that is only revealed after cleavage and release of the MUC1 tandem repeat domain. Neither huMNC2-CAR44 nor huMNC2-CAR50 T cells recognize the cells expressing full-length MUC1 (Fig. 45F-45H). However, when MMP9 plus activator APMA is added, the CAR T cells recognize the cells, cluster and kill them (Fig. 45J-45L). The addition of cleavage enzyme ADAM-17 did not affect the recognition of either CAR T cell for full-length MUC1 (Fig. 45N-45P). The reason could be that ADAM-17 doesn’t cleave MUC1 or the cleavage product is not recognized by MNC2. A similar experiment was performed (Fig. 46) that showed that MMP2 was only weakly effective at either cleavage MUC1 or that the MMP2

cleavage product was only weakly recognized by MNC2. Figure 47 shows the contrast between huMNC2-CAR44 recognition of HCT-MUC1\* cells, T47D-wt breast cancer cells, and T47D cells with added MMP9 which presumably cleaves the full-length MUC1 to an MNC2 recognizable MUC1\*.

**[00770] 3. Confocal imaging of CAR T cells giving the “kiss of death” to MUC1\* positive cancer cells.**

**[00771]** Confocal images of Human T cells that were transduced with huMNC2-CAR44, co-cultured for 24 hours with MUC1\* positive DU145 prostate cancer cells showed the CAR T cells inserting Granzyme B into the target cancer cells. Figure 55 shows fluorescent images of the huMNC2-CAR44 T cells secreting Granzyme B when co-cultured with the prostate cancer cells, FACS analysis showing increased expression of Granzyme B by the CAR T cells and an xCELLigence experiment showing that the target prostate cancer cells were in fact killed.

**[00772] 5. Analysis of CAR T cell induced killing of MUC1\* positive cancer cells by FACS analysis**

**[00773]** We have demonstrated the killing effect of huMNC2-CAR44 T cells on T47D MUC1\* positive breast cancer cells, wherein the breast cancer cells have been transfected with increasing amounts of additional MUC1\*. The killing effect of the huMNC2-CAR44 T cells increases as the amount of target MUC1\* expressed on the cells increases.

**[00774]** IFN- $\gamma$  secretion in media was measured using a human IFN- $\gamma$  ELISA kit (Biolegend). Plates were coated with an anti- IFN- $\gamma$  antibody (capture antibody, 1X in coating buffer). After overnight incubation at 4°C, the plate was washed 4 times with PBS-T and blocking solution was added to block remaining binding site on the well. After 1h at RT (shaking at 500rpm) the plate was washed 4 times with PBS-T and conditioned media (CM) and IFN- $\gamma$  standard, was added. After 2h at RT with shaking, the plate was washed 4 times with PBS-T and detection antibody (1x), was added. After 1h at RT with shaking, the plate was washed 4 times with PBS-T and Avidin-HRP (1x) was added. After 30min at RT with shaking, the plate was washed 5 times with PBS-T (soak 1min each wash) and TMB substrate solution was added. The reaction was stopped after 20min by adding the stop solution and absorbance was read at 450nm (minus absorbance at 570nm) within 15 min of stopping.

**[00775] 6. Analysis of CAR T cell induced killing of MUC1\* positive cancer cells by xCELLigence**

[00776] In addition to FACS analysis, many researchers now use an xCELLigence instrument to measure CAR T killing of cancer cells. The xCELLigence instrument uses electrode arrays upon which cancer cells are plated. The adherent cancer cells insulate the electrode and so cause an increase in impedance as they grow. Conversely, T cells are not adherent and remain in suspension so do not contribute to insulation of the electrode which would increase impedance. However, if the T cells or CAR T cells kill the cancer cells on the electrode plate, the cancer cells ball up and float off as they die, which causes the impedance to decrease. The xCELLigence instrument measures impedance as a function of time, which is correlated to cancer cell killing. In addition, the electrode plates also have a viewing window. When CAR T cells effectively kill the adsorbed target cancer cells, there is a decrease in impedance but also one can see that there are no cancer cells left on the plate surface.

[00777] In most of the XCELLigence experiments, 5,000 cancer cells were plated per well of a 96-well electrode array plate. Cells were allowed to adhere and grow for 24 hours. CAR T cells were then added at an Effector to Target ratio (E:T) of 0.5:1, 1:1, 2:1, 5:1, 10:1 and sometimes 20:1. The E:T ratio assumes 100% transduction of the CAR into the T cells, when the actual transduction efficiency is 40%.

[00778] The xCELLigence instrument records impedance as a function of time and experiments can go on for up to 7 days.

[00779] **Fig. 48, Fig. 49, Fig. 55H, Fig. 56H, Figs. 57A-57C**, all show results of CAR T and cancer cell experiments performed on an xCELLigence instrument.

[00780] **7. Anti-MUC1\* CAR T cell therapy in mice bearing human tumors**

[00781] Female NOD/SCID/GAMMA (NSG) mice between 8-12 weeks of age were implanted with 500,000 human cancer cells, wherein the cancer cells had previously been stably transfected with Luciferase. Mice bearing Luciferase positive cells can be injected with the enzyme's substrate Luciferin just prior to imaging, which makes the cancer cells fluoresce. The cancer cells are imaged in live mice within 10-15 minutes after injection with Luciferin on an IVIS instrument. The readout is flux or photons per second. Tumors were allowed to engraft until tumors were clearly visible by IVIS.

[00782] **Figures 58A-58F** show fluorescent photographs of mice taken on an IVIS instrument. 10 minutes prior to IVIS photographs, mice were injected intraperitoneally (IP) with Luciferin, which fluoresces after cleavage by Luciferase, thus making tumor cells fluoresce. NSG

(NOD/SCID/GAMMA) immune compromised mice that on Day 0 were subcutaneously implanted on the flank with 500,000 human MUC1\* positive cancer cells that had been stably transfected with Luciferase. Tumors were allowed to engraft. On Day 7 after IVIS measurement, animals were tail vein injected with either PBS, 10 million untransduced human T cells or 8.5M huMNC2-scFv-CAR44 T cells. As can be seen in the figure, control mice had to be sacrificed on Day 20 due to excess tumor burden (Fig. 58A-58B). huMNC2-CAR44 T cell treated mice were tumor free after a single CAR T cell injection until Day 100 when they were sacrificed (Fig. 58C). Figure 58E shows Kaplan-Meier survival curves that demonstrate the efficacy of T cell therapy guided by anti-MUC1\* antibody. Figure 58F shows a table summarizing the characteristics of the human T cells that were collected from the test mice upon sacrifice. The starting Car T cell population was 50% CD4 positive helper T cells and 50% CD8 positive killer T cells. As can be seen in the table, the percent of CD8 positive cells has increased in the CAR T treated group, indicating in vivo expansion of that group of cells, which is an indicator of efficacy. We also note that in the treated group, the CAR T cells express higher levels of PD1 which is a marker of T cell exhaustion.

**[00783]** In another animal experiment, NSG mice were sub-cutaneously implanted into the flank with 500,000 tumor cells then injected on Day 7 and again on Day 14 with either saline solution, PBS, or 10M huMNC2-CAR44 T cells (Fig. 59A-59C). In this experiment the amount of MUC1\* expressed on the tumor cells was varied. In one case, the tumor cells that were implanted were T47D-wildtype (Fig. 59B). In another case, the T47D cells were doped with 95% T47D cells that had been transfected to express even more MUC1\* (Fig. 59C). As can be seen, the tumors comprised of cells expressing more MUC1\* were eliminated more quickly and did not recur. In a similar experiment, the tumor cells were doped with a relatively small amount of cells that expressed more MUC1\*. Figure 60A-60C shows NSG mice implanted with T47D-wt breast cancer cells that have been doped with 30% of T47D cells transfected to express more MUC1\*. As can be seen, even a small percentage of cells expressing high levels of MUC1\* is sufficient to trigger CAR T cell mediated killing of the entire tumor. Naturally occurring tumors are heterogeneous and are comprised of both high and low antigen expressing cells. This experiment indicates that huMNC2-CAR44 T cells would be effective in eradicating naturally occurring tumors.

**[00784] Figures 61A-61J** show fluorescent photographs of mice taken on an IVIS instrument. NSG (NOD/SCID/GAMMA) immune compromised mice that on Day 0 were subcutaneously injected into the flank with 500K human BT-20 cells which are a MUC1\* positive triple negative breast cancer cell line. The cancer cells had been stably transfected with Luciferase. Tumors were allowed to engraft. On Day 6 after IVIS measurement, animals were given a one-time injection of 10 million of either human T cells transduced with huMNC2-scFv-CAR44 or untransduced T cells. 5 million T cells were injected intra-tumor and 5 million were injected into the tail vein. 10 minutes prior to IVIS photographs, mice were IP injected with Luciferin. In one case the huMNC2-CAR44 T cells were first incubated with beads to which was attached the PSMGFR peptide to pre-stimulate the T cells and in the figure is marked Protocol 1. In Protocol 2, the huMNC2-CAR44 T cells were pre-stimulated with live tumor cells, which likely injected more tumor cells into the animals' circulation.

**[00785] Figures 62A-62M** show fluorescent photographs of mice taken on an IVIS instrument. NSG (NOD/SCID/GAMMA) immune compromised mice that on Day 0 were injected into the intraperitoneal cavity (IP) with 500K human SKOV-3 cells which are a MUC1\* positive ovarian cancer cell line. The cancer cells had been stably transfected with Luciferase. Tumors were allowed to engraft. On Day 3 after IVIS measurement, animals were IP injected with 10M either human T cells transduced with huMNC2-CAR44 T cells, untransduced T cells or PBS. Animals were IVIS imaged again on Day 7. 10 minutes prior to IVIS photographs, mice were IP injected with Luciferin. As can be seen in the figure the anti-MUC1\* CAR T cells effectively reduced ovarian tumor volume by Day 15.

**[00786] 9. NFAT-induced IL-18 sequences and cloning**

**[00787]** Cloning of IL18 in pGL4-14 3xNFAT:

**[00788]** An activated IL18 (SEQ ID NO:1644) was synthesized with the CD8 leader sequence. The pGL4-14 3xIL2 NFAT and pGL4-14 3xFoxP3 NFAT were digested with XhoI and HindIII restriction enzymes (New England Biolabs). The purified plasmids and the synthesized IL18 sequences were assembled using the Gibson assembly cloning kit (New England Biolab). The resulting constructs (pGL4-14 3xIL2NFAT-IL18 and pGL4-14 3xFoxP3NFAT-IL18) contains 3 repeats of NFAT response element (IL2 or FoxP3) followed by a minimum promoter (mCMV: SEQ ID NO:1634) and IL18 (SEQ ID NOS:1752-1753) with CD8 leader sequence.

**[00789] Cloning of MNC2 CAR with IL18 in pCDNA vector:**

**[00790]** MNC2 CAR sequence was amplified from previously made vector by polymerase chain reaction (PCR) using the following primers: 5'-aggagacccaagctggctagttaagcttgatggccttaccagtgaccgccttgc-3' (SEQ ID NO:1754) and 5'-taggccagagaaatgttctggcattatcagcgagggggcagggcctgc-3' (SEQ ID NO:1755).

**[00791]** IL18 sequence including NFAT response element was amplified from pGL4-14 3xNFAT-IL18 by polymerase chain reaction (PCR) using the following primers: 5'-tgccagaacatttctctgg-3' (SEQ ID NO:1756) and 5'-acagtcgaggctgatcagcggtttaacttatcagtcctcgttctgcacgg-3' (SEQ ID NO: 1757). The purified PCR fragments and digested pCDNA 3.1 V5 (ThermoFisher scientific) were assembled using the Gibson assembly cloning kit (New England Biolab) to create the construct pCDNA MNC2CAR-3xIL2NFAT-IL18 and pCDNA MNC2CAR-3xFoxP3NFAT-IL18.

**[00792] Cloning of MNC2 CAR-NFAT-IL18 in lentivector:**

**[00793]** MNC2 CAR-NFAT-IL18 sequence was amplified from pCDNA MNC2CAR-3xIL2NFAT-IL18 and pCDNA MNC2CAR-3xFoxP3NFAT-IL18 by polymerase chain reaction (PCR) using the following primers: 5'-atgcaggccctgccccctcgctgataagtttaactgccagaacatttctctggcctaac-3' (SEQ ID NO:1758) and 5'-accggagcgatcgagatccttcgcgccgcttatcagtcctcgttctgcacggtgaac-3' (SEQ ID NO:1759). The purified PCR fragments and digested pCDH Dual Hygro (System Biosciences, CA) were assembled using the Gibson assembly cloning kit (New England Biolab) to create the construct pCDH MNC2CAR-3xIL2NFAT-IL18 and pCDH MNC2CAR-3xFoxP3NFAT-IL18.

**[00794] Creation of lentivector with MSCV promoter**

**[00795]** MSCV promoter sequence was amplified from pCDH-MSCV-MCS-EF1a-GFP (System Biosciences) by polymerase chain reaction (PCR) using the following primers: 5'-attgcactagttgaaagacccacctgtagg-3' (SEQ ID NO:1760) and 5'-aatgctctagaatacgggtatccagg-3' (SEQ ID NO:1761). After digestion with SpeI and XbaI restriction enzymes (New England Biolabs), the purified fragment was cloned into pCDH CMV MCS (System Bioscience) digested with the same restriction enzymes to create the construct pCDH MSCV MCS.

**[00796] Cloning of MNC2 CAR-NFAT-IL18 in pCDH MSCV MCS:**

**[00797]** MNC2 CAR-IL2NFAT-IL18 sequence was amplified from pCDNA MNC2CAR-3xIL2NFAT-IL18 by polymerase chain reaction (PCR) using the following primers: 5'

atagcgaattcgtagccgagggccaccatgg-3' (SEQ ID NO:1762) and 5'-taggcctcccaccgtacacgcctaggtaccacgccttctgtatg-3' (SEQ ID NO:1763) MNC2 CAR-IL2NFAT-IL18 sequence was amplified from pCDNA MNC2CAR-3xFoxP3NFAT-IL18 by polymerase chain reaction (PCR) using the following primers: 5' atagcgaattcgtagccgagggccaccatgg -3' (SEQ ID NO:1762) and 5'- taggcctcccaccgtacacgcctaggtacacctgcagtaaattg-3' (SEQ ID NO:1764). After digestion with EcoRI and KpnI restriction enzymes (New England Biolabs), the purified fragment was cloned into pCDH MSCV MCS digested with the same restriction enzymes to create the construct pCDH MSCV MNC2CAR-3xIL2NFAT-IL18 and pCDH MSCV MNC2CAR-3xFoxP3NFAT-IL18.

**[00798] Cloning of 6xNFAT response elements:**

**[00799]** 6xNFAT (IL2 and FoxP3) response element were synthesized followed by different minimal promoter: mCMV (SEQ ID NO:1634), mIL2P (SEQ ID NO:1635) and miniP (SEQ ID NO:1636). A total of six 6 sequences were synthesized: SEQ ID NOS: 1768-1779.

**[00800]** 6xNFAT sequences were amplified by polymerase chain reaction (PCR) using the following primers: 5'-tgccagaacatttctctgg-3' (SEQ ID NO:1756) and 5'- taaggccatggtggctagc-3' (SEQ ID NO:1765). The purified PCR fragments and digested (KpnI and XhoI) pCDNA MNC2CAR 3XNFAT IL18 were assembled using the Gibson assembly cloning kit (New England Biolab) to create constructs with 6x NFAT response elements in place of the 3x NFAT response elements.

**[00801]** 6xNFAT sequences were amplified, from the pCDNA vector created above, by polymerase chain reaction (PCR) using the following primers: 5'-aataagtttaaactgccagaacatttctctgg-3' (SEQ ID NO:1766) and 5'-atatagcgccgcgttatcagtcctcgttctgcacgg-3' (SEQ ID NO:1767). After digestion with PmeI and NotI restriction enzymes (New England Biolabs), the purified fragments were cloned into pCDH MSCV MNC2CAR digested with the same restriction enzymes to create the construct pCDH MSCV MNC2CAR-6xIL2NFAT-IL18 and pCDH MSCV MNC2CAR-6xFoxP3NFAT-IL18. For each construct 3 minimal promoter were tested.

**[00802] Sequence Listing Free Text**

**[00803]** As regards the use of nucleotide symbols other than a, g, c, t, they follow the convention set forth in WIPO Standard ST.25, Appendix 2, Table 1, wherein k represents t or g; n represents a, c, t or g; m represents a or c;

r represents a or g; s represents c or g; w represents a or t and y represents c or t.

#### **MUC1 Receptor**

(Mucin 1 precursor, Genbank Accession number: P15941)

MTPGTQSPFFLLLLLTVLTVVTGSGHASSTPGGEKETSATQRSSVPSSTEKNAVSMSTSSVLSSHSPGSGSSTTQGGQDV  
TLAPATEPASGSAATWGQDVTSVPVTRPALGSTTPPAHDVTSAPDNKPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAH  
GVTSAPDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPP  
AHGVTSAPDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTA  
PPAHGVTSAPDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGS  
TAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAP  
GSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRP  
APGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTR  
RPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA P  
DTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVT  
APDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHG  
TSAPDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAHGVTSA PDTRPAPGSTAPPAH  
GVTSAPDNRPALGSTAPPVHNVTASGSASGSASTLVHNGTSARATTTT PASKSTPFSIPSHHSDTPTTLASHSTKTDA  
SSTHHSSVPPLTSSNHSTSPQLSTGVSTGVSFFFLSFHISNLQFNSSLED PSTDYQELQDISSEMFLQIYKQGGFLGLSNI  
KFRPGSVVVQLTLAFREGTINVHDVETQFNQYKTEAASRYNLTISDVSVDVFPFSAQSGAGVPGWGIALLVLCVL  
VALAIVYLIALAVCQCRRKNYGLDIFPARDTYHPMSEYPTYHTHGRYVPPSSDRSPYEKVSAGNGGSSLSYTNPAV  
AAASANL (SEQ ID NO:1)

#### **PSMGFR**

GTINVHDVETQFNQYKTEAASRYNLTISDVSVDVFPFSAQSGA (SEQ ID NO: 2)

**[00804] N-10 peptide**

**[00805] QFNQYKTEAASRYNLTISDVSVDVFPFSAQSGA (SEQ ID NO:3)**

**[00806] N-19**

**[00807] ASRYNLTISDVSVDVFPFSAQSGA (SEQ ID NO:4)**

**[00808] N-23**

**[00809] NLTISDVSVDVFPFSAQSGA (SEQ ID NO:5)**

**[00810] N-26**

**[00811] ISDVSVDVFPFSAQSGA (SEQ ID NO:6)**

**[00812] N-30**

**[00813] SVSDVFPFSAQSGA (SEQ ID NO:7)**

**[00814] N-10/C-5**

**[00815] QFNQYKTEAASRYNLTISDVSVDVFPFSAQSGA (SEQ ID NO:8)**

**[00816] N-19/C-5**

**[00817] ASRYNLTISDVSVDVFPFSAQSGA (SEQ ID NO:9)**

**[00818] N-36**

**[00819] FPFSAQSGA (SEQ ID NO:10)**

#### **Mouse E6 Heavy chain variable region sequence:**

(DNA)



gaggtgaaggtggtggagtctgggggagacttagtgaagcctggaggggtccctgaaactctcctgtgtagtctctgga  
ttcactttcagtagatatggcatgtcttgggttcgccagactccaggcaagaggctggagtgggtcgcaaccattagt  
ggtggcggtacttacatctactatccagacagtgtgaagggcgattcaccatctccagagacaatgccagaacacc  
ctgtacctgcaaatgagcagtctgaagtctgaggacacagccatgtatcactgtacaagggataactacggtaggaac  
tacgactacggtatggactactgggggtcaaggaacctcagtcaccgtctcctca (SEQ ID NO:12)

(amino acids)

EVKVVESGGDLVKPGGSLKLSCVVS GFTFSRYGMSWVRQTPGKRLEWVATISGGGTYYYPDSVKGRFTISRDNANT  
LYLQMSLLKSEDTAMVHCTRDNYGRNYDYGMDYWGQTSVTVSS (SEQ ID NO:13)

**Mouse E6 heavy chain variable framework region 1 (FWR1) sequence:**

(DNA)

gaggtgaaggtggtggagtctgggggagacttagtgaagcctggaggggtccctgaaactctcctgtgtagtctct  
(SEQ ID NO:14)

(amino acids)

EVKVVESGGDLVKPGGSLKLSCVVS GFTFS (SEQ ID NO:15)

**Mouse E6 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

ggattcactttcagtagatatggcatgtct (SEQ ID NO:16)

(amino acids)

RYGMS (SEQ ID NO:17)

**Mouse E6 heavy chain variable framework region 2 (FWR2) sequence:**

(DNA)

tgggttcgccagactccaggcaagaggctggagtgggtcgca (SEQ ID NO:18)

(amino acids)

WVRQTPGKRLEWVA (SEQ ID NO:19)

**Mouse E6 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

accattagtgggtggcggtacttacatctactatccagacagtgtgaagggg (SEQ ID NO:20)

(amino acids)

TISGGGTYYYPDSVKG (SEQ ID NO:21)

**Mouse E6 heavy chain variable framework region 3 (FWR3) acid sequence:**

(DNA)

cgattcaccatctccagagacaatgccagaacacctgtacctgcaaatgagcagtctgaagtctgaggacacagc  
catgtatcactgtacaagg (SEQ ID NO:22)

(amino acids)

RFTISRDNANTLYLQMSLLKSEDTAMVHCTR (SEQ ID NO:23)

**Mouse E6 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

gataactacggtaggaactacgactacggtatggactac (SEQ ID NO:24)

(amino acids)

DNYGRNYDYGMDY (SEQ ID NO:25)

**Humanized E6 heavy chain variable region sequence from IGHV3-21\*03:**

(DNA)

gaggtgcagctggtggagctctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgg  
attcaccttcagtaggtatggcatgagctgggtccgccaggctccaggaagaggctggagtgggtctcaaccatta  
gtggcggaggcacctacatatactaccagactcagtgaagggccgattcaccatctccagagacaacgccaagaac  
accctgtatctgcaaataaacagcctgagagccgaggacacggctgtgtattactgtaccagagataactatggccg  
caactatgattatggcatggattattggggccagggcaccctggtgaccgtgagcagc (SEQ ID NO:38)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGTYYYPDSVKGRFTISRDNANKN  
TLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGTILVTVSS (SEQ ID NO:39)

**Humanized E6 heavy chain variable framework region 1 (FWR1) acid sequence:**

(DNA)

gaggtgcagctggtggagctctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgg  
attcaccttcagt (SEQ ID NO:40)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFS (SEQ ID NO:41)

**Humanized E6 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

aggtatggcatgagc (SEQ ID NO:42)

(amino acids)

RYGMS (SEQ ID NO:43)

**Humanized E6 heavy chain variable framework region 2 (FWR2) acid sequence:**

(DNA)

tgggtccgccaggctccaggaagaggctggagtgggtctca (SEQ ID NO:44)

(amino acids)

WVRQAPGKRLEWVS (SEQ ID NO:45)

**Humanized E6 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

accattagtggcggaggcacctacatatactaccagactcagtgaagggc (SEQ ID NO:46)

(amino acids)

TISGGGTYYYPDSVKG (SEQ ID NO:47)

**Humanized E6 heavy chain variable framework region 3 (FWR3) acid sequence:**

(DNA)

cgattcaccatctccagagacaacgccaagaacaccctgtatctgcaaataaacagcctgagagccgaggacacggct  
gtgtattactgtaccaga (SEQ ID NO:48)

(amino acids)

RFTISRDNANKNTLYLQMNSLRAEDTAVYYCTR (SEQ ID NO:49)

**Humanized E6 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

gataactatggccgcaactatgattatggcatggattat (SEQ ID NO:50)

(amino acids)

DNYGRNYDYGMDY (SEQ ID NO:51)

**Humanized E6 IgG2 heavy chain synthesized by Genescript:**

(DNA)

gaattctaagcttggggccaccatggaactggggctccgctgggttttcttctgttctattttagaaggtgtccagtgt  
 gaggtgcagctggtggagtctgggggaggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagtaggtatggcatgagctgggtccgccagggtccagggaagaggtgaggtgggtctcaaccattagt  
 ggcgagggcacctacatatactaccagactcagtggaagggccgattcaccatctccagagacaacgccaaagacacc  
 ctgtatctgcaaatgaacagcctgagagccgaggacacggctgtgtattactgtaccagagataactatggccgcaac  
 tatgattatggcatggattattggggccagggcaccctgggtgacgctgagcagcgcctccaccaaggggcccatcggtc  
 tccccctgggcgcctgtccaggagcacctccgagagcacagcgcgcctgggtgctggtcaaggactacttcccc  
 gaaccggtgacggtgtcgtggaactcaggcgtctgtaccagcggcgtgcacacctcccagctgtcctacagtcctca  
 ggactctactccctcagcagcgtggtgaccgtgccctccagcaacttcggcaccagacctacacctgcaacgtagat  
 cacaagcccagcaacaccaaggtggacaagacagttgagcgcgaatgttgtgtcagtgcccaccgtgcccagcacca  
 cctgtggcaggaccgtcagtccttctcttccccccaaaacccaaggacacctcatgatctcccggaacctgaggtc  
 acgtgcgtggtggtggagcgtgagccacgaagaccccgaggtccagttcaactggtacgtggacggcgtggaggtgcat  
 aatgccaagacaaagccacgggaggagcagttcaacagcaggttccgtgtggtcagcgtcctcaccgttgtgcaccag  
 gactggctgaacggcaaggagtacaagtgaaggtctccaacaaaggcctccagcccccatcgagaaaaccatctcc  
 aaaaccaaagggcagccccgagaaccacaggtgtacacctgcccccatcccgaggagatgaccaagaaccaggtc  
 agcctgacctgctggtcaaaggcttctaccccagcgacatcgccgtggagtgggagagcaatgggcagccggagaac  
 aactacaagaccacacctcccatgctggactccgacggctccttcttctctacagcaagctcaccgtggacaagagc  
 aggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggtctctgcacaaccactacacgcagaagagcctc  
 tcctgtctccgggtaaatagtaagtttaactctaga (SEQ ID NO:52)

(amino acids)

EF\*AWATMELGLRWVFLVAILEGVQCEVQLVESGGGLVKPGGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTIS  
 GGGTYIYYPDSVKGRFTISRDNKNTLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLVTVSSASTKGPSV  
 FPLAPCSRSTSESTAALGCLVKDYFPEPVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVTVPSNFGTQTYTCNVD  
 HKPSNTKVDKTVKCCVECPPCAPPVAGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVQFNWYVDGVEVH  
 NAKTKPREEQFNSTFRVSVLTVVHQDWLNGKEYKCKVSNKGLPAPIEKTISKTKGQPREPQVYTLPPSREEMTKNQV  
 SLTCLVKGFIYPSDIAVEWESNGQPENNYKTTTPMLDSDGSFFLYSKLTVDKSRWQQGNVFCSSVMHEALHNHYTQKSL  
 SLSPGK\*\*V\*TLX (SEQ ID NO:53)

**Human IgG2 heavy chain constant region sequence:**

(DNA)

gcctccaccaaggggcccatcggtcttccccctgggcgcctgtctccaggagcacctccgagagcacagcgcgcctgggc  
 tgctggtcaaggactacttccccgaaccggtgacggtgtcgtggaactcaggcgtctgtaccagcggcgtgcacacc  
 tcccagctgtcctacagtcctcaggactctactccctcagcagcgtggtgacgtgccctccagcaacttcggcacc  
 cagacctacacctgcaacgtagatcacaagcccagcaacaccaaggtggacaagacagttgagcgcgaatgttgtgtc  
 gagtgtcccaccgtgcccagcaccacctgtggcaggaccgtcagtccttctcttccccccaaaacccaaggacacctc  
 atgatctcccggaacctgaggtcagtgctggtggtggacgtgagccacgaagaccccgaggtccagttcaactgg  
 tacgtggacggcgtggaggtgcataatgccaagacaaagccacgggaggagcagttcaacagcaggtccgtgtggtc  
 agcgtcctcaccgttgtgtcaccaggactggctgaacggcaaggagtacaagtgaaggtctccaacaaaggcctccca  
 gcccccatcgagaaaaccatctccaaaaccaaagggcagccccgagaaccacaggtgtacacctgcccccatcccg  
 gaggagatgaccaagaaccaggtcagcctgacctgctggtcaaaggcttctaccccagcgacatcgccgtggagtgg  
 gagagcaatgggcagccggagaacaactacaagaccacacctcccatgctggactccgacggctccttcttctctac  
 agcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggtctctgcac  
 aaccactacacgcagaagagcctctcctgtctccgggtaaatag (SEQ ID NO:54)

(amino acids)

ASTKGPSVFPLAPCSRSTSESTAALGCLVKDYFPEPVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVTVPSNFGT  
 QTYTCNVDHKPSNTKVDKTVKCCVECPPCAPPVAGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVQFNW  
 YVDGVEVHNKTKPREEQFNSTFRVSVLTVVHQDWLNGKEYKCKVSNKGLPAPIEKTISKTKGQPREPQVYTLPPSR  
 EEMTKNQVSLTCLVKGFIYPSDIAVEWESNGQPENNYKTTTPMLDSDGSFFLYSKLTVDKSRWQQGNVFCSSVMHEALH  
 NHYTQKSLSLSPGK (SEQ ID NO:55)

**Humanized E6 IgG1 heavy chain sequence:**

(DNA)

gaggtgcagctggtggagctctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagtaggtatggcatgagctgggtccgccaggctccagggaagaggctggagtgggtctcaaccattagt  
 ggcgaggacacatactatactaccagactcagtgaagggccgattcaccatctccagagacaacgccaagaaccca  
 ctgtatctgcaaatgaacagcctgagagccgaggacacggctgtgtattactgtcccagagataactatggccgcaac  
 tatgattatggcatggattattggggccagggcaccctggtgaccgtgagcagcgttagcaccaagggcccatcggtc  
 tccccctggcaccctcctccaagagcacctctgggggcacagcgccctgggtgacctggtcaaggactacttcccc  
 gaaccggtgacggtgtcgtggaactcaggcgccctgaccagcggtgcacacctcccggtgctcctacagtcctca  
 ggactctactccctcagcagcgtggtgacagtgccctccagcagcttgggcacccagacacatctgcaacgtgaat  
 cacaagcccagcaacaccaaggtggacaagaaagttgagcccaaatcttgtgacaaaactcacacatgccacccgtgc  
 ccagcacctgaactcctggggggacgctcagctcttctcttcccccccaaaacccaaggacacccctcatgatctcccg  
 accctgaggtcacatgctggtggtggagctgagccacgaagaccctgaggtcaagttcaactggtacgtggacggc  
 gtggaggtgcataatgccaagacaaagccgaggaggagcagtagacaacagcagctaccgtgtggtcagcgtcctcacc  
 gtctgacaccaggactggctgaatggcaaggagtacaagtgaaggtctccaacaaagccctcccagcccccatcgag  
 aaaaccatctccaaagccaaagggcagccccgagaaccacaggtgtacacctgcccccatcccgggaggagatgacc  
 aagaaccaggtcagcctgacctggtggtcaaaggcttctatcccagcgacatcgccgtggagtgggagagcaatggg  
 cagccggagaacaactacaagaccacgcctcccgctgctggactccgacggctccttcttctctacagcaagctcacc  
 gtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggtcttgcaacaaccactacacg  
 cagaagagcctctccctgtctccgggtaaatgataa (SEQ ID NO:56)

(amino acids)

EVQLVESGGGLVPGGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGTYYIYPDSVKGRFTISRDNKPNP  
 LYLQMNSLRAEDTAVYYCPRDNYGRNYDYGMDYWGQGLTVTVSSASTKGPSVFPLAPSSKSTSGGTAALGCLVKDYFP  
 EPVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVTVPSSSLGTQTYICNVNHKPSNTKVDKKVEPKSCDKTHTCPPC  
 PAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVKFNWYVDGVEVHNAKTKPREEQYNSTYRVVSVLT  
 VLHQDWLNGKEYKCKVSNKALPAPIEKTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNG  
 QPENNYKTTTPVLDSGDSFFLYSKLTVDKSRWQQGNVSCSVMHEALHNHYTQKSLSLSPGK\*\* (SEQ ID NO:57)

#### Human IgG1 heavy chain constant region sequence:

(DNA)

gctagcaccagggccccatcggtcttccccctggcaccctcctccaagagcacctctgggggcacagcgccctgggc  
 tgctggtcaaggactacttccccgaaccggtgacggtgtcgtggaactcaggcgccctgaccagcggtgcacacc  
 tccccggtgtcctacagtcctcaggactctactccctcagcagcgtggtgacagtgcctccagcagcttgggcacc  
 cagacctacatctgcaacgtgaatcacaagcccagcaacaccaaggtggacaagaaagttgagcccaaatcttgtgac  
 aaaactcacacatgccacccgtgccagcacctgaactcctggggggacgctcagctcttctcttcccccccaaaaccc  
 aaggacacccctcatgatctcccgacccctgaggtcacatgctggtggtggacgtgagccacgaagaccctgaggtc  
 aagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagccgaggaggagcagtagacaacagcacg  
 taccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggagtacaagtgaaggtctccaac  
 aaagccctcccagcccccatcgagaaaaccatctccaagccaaagggcagccccgagaaccacaggtgtacacctg  
 cccccatcccgggaggagatgaccaagaaccaggtcagcctgacctgctggtcaaaggcttctatcccagcgacatc  
 gccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgctgctggactccgacggctcc  
 ttcttctctacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcat  
 gaggtcttgcaacaaccactacacgcagaagagcctctccctgtctccgggtaaatgataa (SEQ ID NO:58)

(amino acids)

ASTKGPSVFPLAPSSKSTSGGTAALGCLVKDYFPEPVTWSNSGALTSGVHTFPAVLQSSGLYSLSSVTVPSSSLGT  
 QTYICNVNHKPSNTKVDKKVEPKSCDKTHTCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEV  
 KFNWYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAKGQPREPQVYTL  
 PPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTTPVLDSGDSFFLYSKLTVDKSRWQQGNVSCSVMH  
 EALHNHYTQKSLSLSPGK\*\* (SEQ ID NO:59)

#### Mouse E6 Light Chain variable region sequence:

(DNA)

caaattgttctcaccagctctccagcaatcatgtctgcatctccaggggaggaggtcacccaaacctgcagtgccacc  
 tcaagtgttaagttacatacactggttccagcagaggccaggcaattctcccaaacctctggatttatagcacatccaac  
 ctggcttctggagtccctgttcgcttcagtggcagtggtatgggacctcttactctctcacaatcagccgaatggag

gctgaagatgctgccacttattactgccagcaaaggagtagttccccattcacgttcggctcggggacaaagttggaa  
 ataaaa (SEQ ID NO:65)  
 (amino acids)  
 QIVLTQSPAIMASASPGEVTLTCSATSSVSYIHWFQQRPGTSPKLWIYSTSNLASGVPVRFSGSGYGTSYSLTISRME  
 AEDAATYYCQQRSSSPFTFGSGTKLEIK (SEQ ID NO:66)

**Mouse E6 light chain variable framework region 1 (FWR1) sequence:**

(DNA)  
 caaattgttctcaccagtcctccagcaatcatgtctgcattctccagggaggaggtcacctaacctgc (SEQ ID  
 NO:67)

(amino acids)  
 QIVLTQSPAIMASASPGEVTLTC (SEQ ID NO:68)

**Mouse E6 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)  
 AGTGCCACCTCAAGTGTAAGTTACATACAC (SEQ ID NO:69)

(amino acids)  
 SATSSVSYIH (SEQ ID NO:70)

**Mouse E6 light chain variable framework region 2 (FWR2) sequence:**

(DNA)  
 tggttccagcagaggccaggcacttctcccaaactctggatttat (SEQ ID NO:71)

(amino acids)  
 WFQQRPGTSPKLWIY (SEQ ID NO:72)

**Mouse E6 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)  
 agcacatccaacctggcttct (SEQ ID NO:73)

(amino acids)  
 STSNLAS (SEQ ID NO:74)

**Mouse E6 light chain variable framework region 3 (FWR3) sequence:**

(DNA)  
 ggagtccttggttcgcttcagtgccagtgatattgggacctcttactctctcacaaatcagccgaatggaggctgaaga  
 tgctgccacttattactgc (SEQ ID NO:75)

(amino acids)  
 GVPVRFSGSGYGTSYSLTISRMEAEDAATYYC (SEQ ID NO:76)

**Mouse E6 light chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)  
 cagcaaaggagtagttccccattcacg (SEQ ID NO:77)

(amino acids)  
 QQRSSSPFT (SEQ ID NO:78)

**Humanized E6 light chain variable region sequence from IGKV3-11\*02:**

(DNA)  
 gaaattgtgttgacacagtcctccagccacctgtctttgtctccaggggaaagagccacctcacctgcagcgccacc  
 agcagtgtagctacatccactggtaccaacagaggcctggccagagccccagggtcctcatctatagcacctccaac

ctggccagcggcatcccagccaggttcagtggcagtggtctgggagcgactacactctcaccatcagcagcctagag  
cctgaagattttgcagtttattactgtcagcagcgtagcagctccccctttcacctttggcagcggcaccaaagtggaa  
attaaa (SEQ ID NO:93)

(amino acids)

EIVLTQSPATLSLSPGERATLTCSATSSVSYIHQYQQRPGQSPRLLIYSTSNLASGIPARFSGSGSGSDYTLTISSLE  
PEDFAVYYCQQRSSSPFTFGSGTKVEIK (SEQ ID NO:94)

**Humanized E6 light chain variable framework region 1 (FWR1) acid sequence:**

(DNA)

gaaattgtgttgacacagtctccagccaccctgtctttgtctccaggggaaagagccaccctcacctgc (SEQ  
ID NO:95)

(amino acids)

EIVLTQSPATLSLSPGERATLTC (SEQ ID NO:96)

**Humanized E6 light chain variable complementarity determining regions 1  
(CDR1) sequence:**

(DNA)

agcgccaccagcagtggttagctacatccac (SEQ ID NO:97)

(amino acids)

SATSSVSYIH (SEQ ID NO:98)

**Humanized E6 heavy light variable framework region 2 (FWR2) acid sequence:**

(DNA)

tggtaccaacagaggcctggccagagccccaggtctctcatctat (SEQ ID NO:99)

(amino acids)

WYQQRPGQSPRLLIY (SEQ ID NO:100)

**Humanized E6 light chain variable complementarity determining regions 2 (CDR2)  
sequence:**

(DNA)

agcacctccaacctggccagc (SEQ ID NO:101)

(amino acids)

STSNLAS (SEQ ID NO:102)

**Humanized E6 light chain variable framework region 3 (FWR3) acid sequence:**

(DNA)

ggcatcccagccaggttcagtggcagtggtctgggagcgactacactctcaccatcagcagcctagagcctgaagat  
tttgagtttattactgt (SEQ ID NO:103)

(amino acids)

GIPARFSGSGSGSDYTLTISSLEPEDFAVYYC (SEQ ID NO:104)

**Humanized E6 light chain variable complementarity determining regions 3 (CDR3)  
sequence:**

(DNA)

cagcagcgtagcagctccccctttcacc (SEQ ID NO:105)

(amino acids)

QQRSSSPFT (SEQ ID NO:106)

**Humanized E6 Kappa light chain synthesized by Genescript:**

(DNA)

gaattctaagcttgggccaccatggaagccccagcgcagcttctcttctctctgctactctggctcccagataccact  
ggagaaattgtgttgacacagtctccagccaccctgtctttgtctccaggggaaagagccaccctcacctgcagcgcc

accagcagtggttagctacatccactggtaccaacagaggcctggccagagagcccagggtcctcatctatagcacctcc  
aacctggccagcgccatcccagccaggttcagtgccagtggtctgggagcgactacactctcaccatcagcagccta  
gagcctgaagattttgcagtttattactgtcagcagcgtagcagctccccctttcacctttggcagcgccaccaagtg  
gaaattaaaaggacgggtggctgcaccatctgtcttcatcttcccgccatctgatgagcagttgaaatctggaactgcc  
tctgttgtgtgcctgctgaataaacttctatcccagagaggccaaagtacagtgggaaggtggataacgcctccaatcg  
ggtaactcccaggagagtggtcacagagcaggacagcaaggacagcacctacagcctcagcagcacctgacgctgagc  
aaagcagactacgagaaacacaaagtctacgcctgcgaagtcacccatcagggcctgagctcgcccgtcacaaagagc  
ttcaacaggggagagtggttagtaagtttaactctaga (SEQ ID NO:107)

(amino acids)

EF\*AWATMEAPAQLFLLLLWLPD TTGEIVLTQSPATLSLSPGERATLTCSATSSVSYIHWHYQQRPGQSPRLLIYSTS  
NLASGIPARFSGSGSGSDYTLTISSLEPEDFAVYYCQQRSSSPFTFGSGTKVEIKRTVAAPSVFIFPPSDEQLKSGTA  
SVVCLLNNFYPREAKVQWKVDNALQSGNSQESVTEQDSKDYSLSSLTLSKADYEKHKVYACEVTHQGLSSPVTKS  
FNRGEC\*\*V\*TLX (SEQ ID NO:108)

#### Human Kappa light chain constant region sequence:

(DNA)

aggacgggtggctgcaccatctgtcttcatcttcccgccatctgatgagcagttgaaatctggaactgcctctgttgtg  
tgctgctgaataaacttctatcccagagaggccaaagtacagtgggaaggtggataacgcctccaatcggttaactcc  
caggagagtggtcacagagcaggacagcaaggacagcacctacagcctcagcagcacctgacgctgagcaaagcagac  
tacgagaaacacaaagtctacgcctgcgaagtcacccatcagggcctgagctcgcccgtcacaaagagcttcaacagg  
ggagagtggttag (SEQ ID NO:109)

(amino acids)

RTVAAPSVFIFPPSDEQLKSGTASVVCLLNNFYPREAKVQWKVDNALQSGNSQESVTEQDSKDYSLSSLTLSKAD  
YEKHKVYACEVTHQGLSSPVTKSFNRGEC (SEQ ID NO:110)

#### Humanized E6 lambda light chain sequence:

(DNA)

gaaattgtgttgacacagtcctccagccaccctgtctttgtctccaggggaaagagccaccctcacctgcagcgccacc  
agcagtggttagctacatccactggtaccaacagaggcctggccagagagcccagggtcctcatctatagcacctccaac  
ctggccagcgccatcccagccaggttcagtgccagtggtctgggagcgactacactctcaccatcagcagcctagag  
cctgaagattttgcagtttattactgtcagcagcgtagcagctccccctttcacctttggcagcgccaccaagtgga  
attaaaggtcagcccaaggctgccccctcggtcactctgttcccgccctcctctgaggagcttcaagccaacaaggcc  
acactggtgtgtctcataagtgaattctacccgggagcctgacagtggcctggaaggcagatagcagccccgtcaag  
gcgggagtgaggaccaccacacccctccaaacaaagcaacaacaagtaacggcgccagcagctatctgagcctgacgcct  
gagcagtggaagtcacacagaagctacagctgccaggtcacgcattgaagggagcaccgtggagaagacagtgggccct  
acagaatgttcataagtaa (SEQ ID NO:111)

(amino acids)

EIVLTQSPATLSLSPGERATLTCSATSSVSYIHWHYQQRPGQSPRLLIYSTSNLASGIPARFSGSGSGSDYTLTISSLE  
PEDFAVYYCQQRSSSPFTFGSGTKVEIKGQPKAAPSVTLFPPSSEELQANKATLVCLISDFYPGAVTVAWKADSSPVK  
AGVETTTPSKQSNKYYAASSYLSLTPEQWKSHRSYSCQVTHEGSTVEKTVAPTECS\*\* (SEQ ID NO:112)

#### Humanized lambda light chain constant region sequence:

(DNA)

ggtcagcccaaggctgccccctcggtcactctgttcccgccctcctctgaggagcttcaagccaacaaggccacactg  
gtgtgtctcataagtgaattctacccgggagcctgacagtggcctggaaggcagatagcagccccgtcaaggcgga  
gtggagaccaccacacccctccaaacaaagcaacaacaagtaacggcgccagcagctatctgagcctgacgcctgagcag  
tggaagtcacacagaagctacagctgccaggtcacgcattgaagggagcaccgtggagaagacagtgggccctacagaa  
tgttcatagtaa (SEQ ID NO:113)

(amino acids)

GQPKAAPSVTLFPPSSEELQANKATLVCLISDFYPGAVTVAWKADSSPVKAGVETTTPSKQSNKYYAASSYLSLTPEQ  
WKSHRSYSCQVTHEGSTVEKTVAPTECS\*\* (SEQ ID NO:114)

**Mouse C2 heavy chain variable region sequence:**

(DNA)

gagggtccagctggaggagtcagggggagggttagtgaagcctggaggggtccctgaaactctcctgtgcagcctctgg  
attcactttcagtggtatgccatgtcttgggttcgccagactccggagaagaggctggagtgggtcgcaaccatta  
gtagtgggtggtacttatactactatccagacagtggtgaaggggcgattcaccatctccagagacaatgccaagaac  
accctgtacctgcaaagagcagtcctgaggtctgaggacacggccatgtattactgtgcaagacttgggggggataa  
ttactacgaatacttcgatgtctggggcgaggaccaggtcaccgtctcctccgcaaaaacgacacccccatctg  
tctat (SEQ ID NO:118)

(amino acids)

EVQLEESGGGLVKPGGSLKLSAASGFTFSGYAMSWVRQTPEKRLEWVATISSGGTYIYYPDSVKGRFTISRDNKN  
TLYLQMSSLRSEDAMYYCARLGGDNYEYFDVWGAGTTVTVSSAKTTPPSVY (SEQ ID NO:119)

**Mouse C2 heavy chain variable framework region 1 (FWR1) sequence:**

(DNA)

gagggtccagctggaggagtcagggggagggttagtgaagcctggaggggtccctgaaactctcctgtgcagcctctgg  
attcactttcagt (SEQ ID NO:120)

(amino acids)

EVQLEESGGGLVKPGGSLKLSAASGFTFS (SEQ ID NO:121)

**Mouse C2 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

ggctatgccatgtct (SEQ ID NO:122)

(amino acids)

GYAMS (SEQ ID NO:123)

**Mouse C2 heavy chain variable framework region 2 (FWR2) sequence:**

(DNA)

tgggttcgccagactccggagaagaggctggagtgggtcgca (SEQ ID NO:124)

(amino acids)

WVRQTPEKRLEWVA (SEQ ID NO:125)

**Mouse C2 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

accattagtagtggtggtacttatactactatccagacagtggtgaagggg (SEQ ID NO:126)

(amino acids)

TISSGGTYIYYPDSVKG (SEQ ID NO:127)

**Mouse C2 heavy chain variable framework region 3 (FWR3) sequence:**

(DNA)

cgattcaccatctccagagacaatgccaagaacaccctgtacctgcaaagagcagtcctgaggtctgaggacacggcc  
atgtattactgtgcaaga (SEQ ID NO:128)

(amino acids)

RFTISRDNKN TLYLQMSSLRSEDAMYYCAR (SEQ ID NO:129)

**Mouse C2 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

cttgggggggataattactacgaatacttcgatgtc (SEQ ID NO:130)

(amino acids)



LGGDNYEYFDV (SEQ ID NO:131)

Humanized derived from **IGHV3-21\*04**:

**Humanized C2 heavy chain variable region sequence:**

(DNA)

gaggtgcagctggtggagctctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
ttcaccttcagtggtatgccatgagctgggtccgccaggctccaggaaggggctggagtggtctcaaccattagt  
agtggcggaacctacatatactaccccgactcagtggaagggccgattcaccatctccagagacaacgccagaactca  
ctgtatctgcaaatgaacagcctgagagccgaggacacggccgtgtattactgtgcgagacttgggggggataattac  
tacgaataacttcgatgtctggggcaaagggaccacggtcaccgtctctctcc (SEQ ID NO:144)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSGYAMSWVRQAPGKLEWVSTISSGGTYIYYPDSVKGRFTISRDNKNS  
LYLQMNSLRAEDTAVYYCARLGGDNYEYFDVWGKGTITVTVSS (SEQ ID NO:145)

**Humanized C2 heavy chain variable framework region 1 (FWR1) sequence:**

(DNA)

gaggtgcagctggtggagctctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
ttcaccttcagt (SEQ ID NO:146)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFS (SEQ ID NO:147)

**Humanized C2 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

ggctatgccatgagc (SEQ ID NO:148)

(amino acids)

GYAMS (SEQ ID NO:149)

**Humanized C2 heavy chain variable framework region 2 (FWR2) sequence:**

(DNA)

tgggtccgccaggctccaggaaggggctggagtggtctcaa (SEQ ID NO:150)

(amino acids)

WVRQAPGKLEWVS (SEQ ID NO:151)

**Humanized C2 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

accattagtagtggtcggaacctacatatactaccccgactcagtggaagggc (SEQ ID NO:152)

(amino acids)

TISSGGTYIYYPDSVKG (SEQ ID NO:153)

**Humanized C2 heavy chain variable framework region 3 (FWR3) sequence:**

(DNA)

cgattcaccatctccagagacaacgccagaactcactgtatctgcaaatgaacagcctgagagccgaggacacggcc  
gtgtattactgtgcgaga (SEQ ID NO:154)

(amino acids)

RFTISRDNKNSLYLQMNSLRAEDTAVYYCAR (SEQ ID NO:155)

**Humanized C2 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

cttgggggggataattactacgaataacttcgatgtc (SEQ ID NO:156)

(amino acids)  
 LGGDNYEYFDV (SEQ ID NO:157)

#### Humanized C2 IgG1 heavy chain sequence

(DNA)

gaggtgcagctggtggagctctgggggaggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagtggtatgccatgagctgggtccgccaggctccaggggaaggggctggagtggtctcaaccattagt  
 agtggcggaacctacatatactaccccgactcagtggaagggcgattcaccatctccagagacaacgccagaactca  
 ctgtatctgcaaatgaacagcctgagagccgaggacacggcctgtattactgtgcgagacttgggggggataattac  
 tacgaatacttcgatgtctggggcaaagggaccacgggtcacctctcctccgctagcaccaagggcccatcggtcttc  
 cccctggcaccctcctccaagagcacctctgggggcacagcggcctgggctgctgggtcaaggactacttccccgaa  
 ccggtgacgggtgtcgtggaactcaggcgccctgaccagcggcgtgcacaccttcccggctgtcctacagtcctcagga  
 ctctactccctcagcagcgtggtgacagtgcctccagcagcttgggcacccagacctacatctgcaacgtgaatcac  
 aagcccagcaacaccaaggtggacaagaaagttgagcccaaatcttgtgacaaaactcacacatgccacccgtgcca  
 gcacctgaactcctggggggaccgtcagttcttcttcccccaaaaacccaaggacacctcatgatctcccggacc  
 cctgaggtcacatgctggtggtggacgtgagccacgaagacctgaggtcaagttcaactggtacgtggacggcgtg  
 gaggtgcataatgccaaagacaaagccgaggaggagcagtaaacagcagctaccgtgtggtcagcgtcctcacctgc  
 ctgcaccaggactggctgaatggcaaggagtacaagtgaaggtctccaacaaagccctcccagcccccatcgagaaa  
 accatctccaaagccaaagggcagccccgagaaccacaggtgtacacctgcccccatcccgggaggagatgaccaag  
 aaccaggtcagcctgacctgctggtcaaaggtctctatcccagcgacatcgccgtggagtgaggagcaatgggcag  
 ccggagaacaactacaagaccacgcctcccgtgctggactccgacggctccttcttctctacagcaagctcacctgc  
 gacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggtctctgcacaaccactacacgcag  
 aagagcctctcctgtctccgggtaaatgataa (SEQ ID NO:157)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGTYIYYPDSVKGRFTISRDNKNS  
 LYLQMNSLR AEDTAVYYCARLGGDNYEYFDVWGKGTITVTVSSASTKGPSVFPLAPSSKSTSGGTAALGCLVKDYFPE  
 PVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVTVTPSSSLGTQTYICNVNHKPSNTKVDKKVEPKSCDKTHTCPPCP  
 APELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHNAKTKPREEQYNSTYRVVSVLTV  
 LHQDWLNGKEYKCKVSNKALPAPIEKTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQ  
 PENNYKTTTPVLDSDGSFFLYSKLTVDKSRWQQGNVSCSVMHEALHNHYTQKSLSLSPGK\*\* (SEQ ID NO:158)

#### Humanized C2 IgG2 heavy chain sequence

(DNA)

gaggtgcagctggtggagctctgggggaggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagtggtatgccatgagctgggtccgccaggctccaggggaaggggctggagtggtctcaaccattagt  
 agtggcggaacctacatatactaccccgactcagtggaagggcgattcaccatctccagagacaacgccagaactca  
 ctgtatctgcaaatgaacagcctgagagccgaggacacggcctgtattactgtgcgagacttgggggggataattac  
 tacgaatacttcgatgtctggggcaaagggaccacgggtcacctctcctccgctccaccaagggcccatcggtcttc  
 cccctggcgccctgctccaggagcacctccgagagcacagcgcctgggctgctgggtcaaggactacttccccgaa  
 ccggtgacgggtgtcgtggaactcaggcgctctgaccagcggcgtgcacaccttcccagctgtcctacagtcctcagga  
 ctctactccctcagcagcgtggtgacctgacctccagcaacttgggcacccagacctacacctgcaacgtagatcac  
 aagcccagcaacaccaaggtggacaagacagttgagcgcgaatgttgtgtcgagtggccacccgtgcccagcaccacct  
 gtggcaggacccgtcagttcttcttcccccaaaaacccaaggacacctcatgatctcccggacccctgaggtcacg  
 tgctggtggtggacgtgagccacgaagacccccgaggtccagttcaactggtacgtggacggcgtggaggtgcataat  
 gccaaagacaaagccacgggaggagcagttcaacagcagttccgtgtggtcagcgtcctcacctgtgtgaccaggac  
 tggctgaacggcaaggagtacaagtgaaggtctccaacaaaggcctcccagcccccatcgagaaaaccatctccaaa  
 accaaagggcagccccgagaaccacaggtgtacacctgcccccatcccgggaggagatgaccaagaaccaggtcagc  
 ctgacctgcctggtcaaaggtcttaccacagcgacatcgccgtggagtgaggagcaatgggcagccggagacaac  
 tacaagaccacacctcccatgctggactccgacggctccttcttctctacagcaagctcacctgggacaagagcagg  
 tggcagcaggggaacgtcttctcatgctccgtgatgcatgaggtctctgcacaaccactacacgcagaagagcctctcc  
 ctgtctccgggtaaatagtaa (SEQ ID NO:163)

(amino acids)

EVQLVESGGGLVQPGGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGTYIYYPD SVKGRFTISRDN AKNS  
 LYLQMNSLR AEDTAVYYCARLG GDNYYEYFDVWGKGTITVTVSSASTKGPSVFPLAPCSRSTSESTAALGCLVKDYFPE  
 PVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVVTVPSSNFGTQTYYTCNV D HKPSNTKVDK TVERKCCVECP P PAPP  
 VAGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVQFNWYVDGVEVHNAKTKPREEQFNSTFRVVS VLT VVH QD  
 WLNGKEYKCKVSNKGLPAPIEKTISKTKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENN  
 YKTTTPMLDS DGSFFLYSKLTVDKSRWQQGNV FSCSV MHEALHNHYTQKSLSLSPGK\*\* (SEQ ID NO:164)

**Mouse C2 light chain variable region sequence:**

(DNA)

gacattgtgatcacacagctctacagcttccttaggtgtatctctggggcagagggccaccatctcatgcagggccagc  
 aaaagtgtcagtacatctggctatagttatatgcactgggtaccaacagagaccaggacagccacccaaactcctcatc  
 tatcttgcatccaacctagaatctggggtccttgccaggttcagtggtcagtggtctgggacagacttcacctcaac  
 atccatcctgtggaggaggaggatgctgcaacctattactgtcagcacagtagggagcttcggttcacgttcggaggg  
 gggaccaagctggagataaaaacgggctgatgctgcaccaactgtatcc (SEQ ID NO:168)

(amino acids)

DIVITQSTASLG VSLGQRATISCRASKSVSTSGYSYMHYQQRPGQPPKLLIYLASNLESGV PARFSGSGSGTDFTLN  
 IHPVEEEDAATYYCQHSREL PFTFGG GTKLEIKRADAAPT VS (SEQ ID NO:169)

**Mouse C2 light chain variable framework region 1 (FWR1) sequence:**

(DNA)

gacattgtgatcacacagctctacagcttccttaggtgtatctctggggcagagggccaccatctcatgc (SEQ ID  
 NO:170)

(amino acids)

DIVITQSTASLG VSLGQRATISC (SEQ ID NO:171)

**Mouse C2 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

agggccagcaaaagtgtcagtacatctggctatagttatatgcac (SEQ ID NO:172)

(amino acids)

RASKSVSTSGYSYMH (SEQ ID NO:173)

**Mouse C2 light chain variable framework region 2 (FWR2) sequence:**

(DNA)

tggtaccaacagagaccaggacagccacccaaactcctcatctat (SEQ ID NO:174)

(amino acids)

WYQQRPGQPPKLLIY (SEQ ID NO:175)

**Mouse C2 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

cttgcatccaacctagaatc (SEQ ID NO:176)

(amino acids)

LASNLES (SEQ ID NO:177)

**Mouse C2 light chain variable framework region 3 (FWR3) sequence:**

(DNA)

tggggtccttgccaggttcagtggtcagtggtctgggacagacttcacctcaacatccatcctgtggaggaggagga  
 tgctgcaacctattactgt (SEQ ID NO:178)

(amino acids)

GVPARFSGSGSGTDFTLNIHPVEEEDAATYYC (SEQ ID NO:179)

**Mouse C2 light chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

cagcacagtagggagcttccggttcacg (SEQ ID NO:180)

(amino acids)

QHSRELPFT (SEQ ID NO:181)

Humanized derived from **IGKV7-3\*01****Humanized C2 light chain variable region sequence:**

(DNA)

gacattgtgctgacccagtcctccagcctccttggccgtgtctccaggacagagggccaccatcacctgcagagccagt  
 aagagtgtcagtagcagcggtactcctacatgcactgggtatcagcagaaaccaggacaacctcctaaactcctgatt  
 tacctggcatccaatctggagagcgggggtccagccaggttcagcggcagtggtctgggaccgatttcacctcaca  
 attaatcctgtggaagctaattgatactgcaaattattactgtcagcacagtagggagctgcctttcacattcggcgga  
 gggaccaaggtggagatcaaacgaact (SEQ ID NO:194)

(amino acids)

DIVLTQSPASLAVSPGQRATITCRASKSVSTSGYSYMHWYQQKPGQPPKLLIYLASNLESGVPARFSGSGSGTDFTLT  
 INPVEANDTANYYCQHSRELPFTFGGGTKVEIKRT (SEQ ID NO:195)

**Humanized C2 light chain variable framework region 1 (FWR1) acid sequence:**

(DNA)

gacattgtgctgacccagtcctccagcctccttggccgtgtctccaggacagagggccaccatcacctgc (SEQ ID NO:196)

(amino acids)

DIVLTQSPASLAVSPGQRATITC (SEQ ID NO:197)

**Humanized C2 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

agagccagtaagagtgtcagtagcagcggtactcctacatgcac (SEQ ID NO:198)

(amino acids)

RASKSVSTSGYSYMH (SEQ ID NO:199)

**Humanized C2 heavy light variable framework region 2 (FWR2) acid sequence:**

(DNA)

tggtatcagcagaaaccaggacaacctcctaaactcctgatttac (SEQ ID NO:200)

(amino acids)

WYQQKPGQPPKLLIY (SEQ ID NO:201)

**Humanized C2 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

ctggcatccaatctggagagc (SEQ ID NO:202)

(amino acids)

LASNLES (SEQ ID NO:203)

**Humanized C2 light chain variable framework region 3 (FWR3) acid sequence:**

(DNA)

gggggtccagccaggttcagcggcagtggtctgggaccgatttcacctcacaattaatcctgtggaagctaattgat  
 actgcaaattattactgt (SEQ ID NO:204)

(amino acids)



ggcggaggcacctacatatactaccagactcagtgaagggcgattcaccatctccagagacaacgccaagaacacc  
ctgtatctgcaaatgaacagcctgagagccgaggacacggctgtgtattactgtaccagagataactatggccgcaac  
tatgattatggcatggattattggggccagggcaccctgggtgaccgtgagcagcgccgggtggcggatccggcggtggc  
ggatccggcggtggcggatccgaaattgtgttgacacagctctccagccaccctgtctttgtctccaggggaaagagcc  
accctcacctgcagcgccaccagcagtgttagctacatccactggtaccaacagaggcctggccagagccccaggctc  
ctcatctatagcacctccaacctggccagcgccatcccagccagggttcagtggcagtggtctgggagcgactacact  
ctcaccatcagcagcctagagcctgaagattttgcagtttattactgtcagcagcgtagcagctccccctttcaccttt  
ggcagcgccaccaaaagtggaaattaaa (SEQ ID NO:232)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGTYYIYPDSVKGRFTISRDN  
AKN TLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLVTVSSGGGSGGGGSGGGGSEIVLTQSPATLSLSPGE  
RATLTCSATSSVSIHWYQQRPGQSPRLLIYSTSNLASGIPARFSGSGSGSDYTLTISLLEPEDFAVYYCQQRSSSP  
FTFGSGTKVEIK (SEQ ID NO:233)

#### Humanized E6 single chain IgG1noC

(DNA)

gaggtgcagctggtggagtctgggggaggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
ttcaccttcagtaggtatggcatgagctgggtccgccaggctccagggaagaggctggagtgggtctcaaccattagt  
ggcggaggcacctacatatactaccagactcagtgaagggcgattcaccatctccagagacaacgccaagaacacc  
ctgtatctgcaaatgaacagcctgagagccgaggacacggctgtgtattactgtaccagagataactatggccgcaac  
tatgattatggcatggattattggggccagggcaccctgggtgaccgtgagcagcgataaaaccataactaaaccgcca  
aaaccggcgccggaactgctgggtgggtcctgggtaccgggtgaaattgtgttgacacagctctccagccaccctgtctttg  
tctccaggggaaagagccaccctcacctgcagcgccaccagcagtgttagctacatccactggtaccaacagaggcct  
ggccagagccccaggctcctcatctatagcacctccaacctggccagcgccatcccagccagggttcagtggcagtggt  
tctgggagcgactacactctcaccatcagcagcctagagcctgaagattttgcagtttattactgtcagcagcgtagc  
agctccccctttcacctttggcagcgccaccaaaagtggaaattaaa (SEQ ID NO:234)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGTYYIYPDSVKGRFTISRDN  
AKN TLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLVTVSSDKTHTKPPKPAPELLGGPGTGEIVLTQSPATL  
SLSPGERATLTCSATSSVSIHWYQQRPGQSPRLLIYSTSNLASGIPARFSGSGSGSDYTLTISLLEPEDFAVYYCQ  
QRSSSPFTFGSGTKVEIK (SEQ ID NO:235)

#### Humanized E6 single chain X4 (linker is IgG1 and IgG2 modified hinge region)

(DNA)

gaggtgcagctggtggagtctgggggaggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
ttcaccttcagtaggtatggcatgagctgggtccgccaggctccagggaagaggctggagtgggtctcaaccattagt  
ggcggaggcacctacatatactaccagactcagtgaagggcgattcaccatctccagagacaacgccaagaacacc  
ctgtatctgcaaatgaacagcctgagagccgaggacacggctgtgtattactgtaccagagataactatggccgcaac  
tatgattatggcatggattattggggccagggcaccctgggtgaccgtgagcagcgataaaaccataactaaaccgcca  
aaaccggcgccggaactgctgggtgggtcctgggtaccgggtactgggtgggtccgactattaaacctccgaaacctccgaaa  
cctgctccgaacctgctgggtgggtccggaattgtgttgacacagctctccagccaccctgtctttgtctccaggggaa  
agagccaccctcacctgcagcgccaccagcagtgttagctacatccactggtaccaacagaggcctggccagagcccc  
aggctcctcatctatagcacctccaacctggccagcgccatcccagccagggttcagtggcagtggtctgggagcgac  
tacactctcaccatcagcagcctagagcctgaagattttgcagtttattactgtcagcagcgtagcagctccccctttc  
acctttggcagcgccaccaaaagtggaaattaaa (SEQ ID NO:236)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGTYYIYPDSVKGRFTISRDN  
AKN TLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLVTVSSDKTHTKPPKPAPELLGGPGTGTGGPTIKPPKP  
PKPAPNLLGGPEIVLTQSPATLSLSPGERATLTCSATSSVSIHWYQQRPGQSPRLLIYSTSNLASGIPARFSGSGS  
GSDYTLTISLLEPEDFAVYYCQQRSSSPFTFGSGTKVEIK (SEQ ID NO:237)

#### Humanized C2 single chain GS3

(DNA)

gaggtgcagctggtggagtctgggggaggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
ttcaccttcagtggtatgccatgagctgggtccgccaggctccagggaaggggctggagtgggtctcaaccattagt  
agtggcggaacctacatatactaccccgactcagtgaagggcgattcaccatctccagagacaacgccaagaactca

ctgtatctgcaaatgaacagcctgagagccgaggacacggccgtgtattactgtgcgagacttgggggggataattac  
tacgaataacttcgatgtctgtggggcaaagggaccacggtcaccgtctcctccggcggtggcggtatccggcggtggcgga  
tccggcggtggcggtatccgacattgtgctgacctcagctctccagcctccttggccgtgtctccaggacagagggccacc  
atcacctgcagagccagtaagagtgtcagtaccagcggatactcctacatgcactgggtatcagcagaaaccaggacaa  
cctcctaaactcctgatttacctggcatccaatctggagagcgggggtcccagccaggttcagcggcagtggtgtggtg  
accgatttcaccctcacaattaatcctgtggaagctaatactgatactgcaaattattactgtcagcacagtagggagctg  
cctttcacattcggcggtggaggaccaaggtggagatcaaacgaact (SEQ ID NO:238)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGTYIYYPDSVKGRFTISRDN  
AKN SLYLQMNSLR AEDTAVYYCARLGDNYYEYFDVWGKGTITVTVSSGGGGSGGGSGGGSDIVLTQSPASLAVSPGQR  
ATITCRASKSVSTSGYSYMHWYQKPGQPPKLLIYLASNLESGVPARFSGSGSGTDFTLTINPVEANDTANYYCQHS  
RELPTFGGGTKVEIKRT (SEQ ID NO:239)

#### Humanized C2 single chain X4 (linker is IgG1 and IgG2 modified hinge region)

(DNA)

gaggtgcagctggtggagtctgtggggaggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
ttcaccttcagtggtctatgccatgagctgggtccgccaggctccagggaaggggtggagtgggtctcaaccattagt  
agtggcggaacctacatatactaccccgactcagtggaagggccgattcaccatctccagagacaacgccagaactca  
ctgtatctgcaaatgaacagcctgagagccgaggacacggccgtgtattactgtgcgagacttgggggggataattac  
tacgaataacttcgatgtctgtggggcaaagggaccacggtcaccgtctcctccgataaaaaccatactaaaccgccaaa  
ccggcgccggaactgctgggtggtcctggtaccggtactggtgggtccgactattaacacctccgaaacctccgaaacct  
gtccgaacctgctgggtggtccggacattgtgctgacctcagctctccagcctccttggccgtgtctccaggacagagg  
gccaccatcacctgcagagccagtaagagtgtcagtaccagcggatactcctacatgcactgggtatcagcagaaacca  
ggacaacctcctaaactcctgatttacctggcatccaatctggagagcgggggtcccagccaggttcagcggcagtggtg  
tctgggaccgatttcaccctcacaattaatcctgtggaagctaatactgatactgcaaattattactgtcagcacagtagg  
gagctgcctttcacattcggcggtggaggaccaaggtggagatcaaacgaact (SEQ ID NO:242)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGTYIYYPDSVKGRFTISRDN  
AKN SLYLQMNSLR AEDTAVYYCARLGDNYYEYFDVWGKGTITVTVSSDKTHTKPPKPAPELLGGPGTGTGGPTIKPPKPP  
KPAPNLLGGPDIVLTQSPASLAVSPGQRATITCRASKSVSTSGYSYMHWYQKPGQPPKLLIYLASNLESGVPARFS  
GSGSGTDFTLTINPVEANDTANYYCQHSRELPTFGGGTKVEIKRT (SEQ ID NO:243)

#### Humanized C3 single chain GS3

(DNA)

caggttcagctggtgcagctctggagctgaggtgaagaagcctggggcctcagtggaaggtctcctgcaaggcttctggt  
tacacctttaccgactacgccatgaactgggtgcgacaggccctggacaagggcttgagtggatgggagtgatcagc  
accttcagcggtaacacaaacttcaaccagaagttcaagggcagagtcaccatgaccacagacacatccacgagcaca  
gcctacatggagctgaggagcctgagatctgacgacacggccgtgtattactgtgcgagaagcgactactacggccca  
tacttcgactactggggccagggcaccacctgaccgtgtccagcggcggtggcggtatccggcggtggcggtatccggc  
ggtggcggtatccgatattgtgatgacctcagactccactctctctgtccgtcaccctggacagccggcctccatctcc  
tgcaggtctagtgcagaccattgtccatagtaatggaacacctatttggagtggtacctgcagaagccaggccagtgct  
ccacagctcctgatctataaggtttccaaccgggtctctgtgagtgccagataggttcagtggcagcgggtcagggaca  
gatttcacactgaaaatcagccgggtggaggctgaggatgttgggggtttattactgcttccaaggttagccacgtgcct  
ttcaccttcggcggtggaggaccaaggtggagatcaaacgaact (SEQ ID NO:244)

(amino acids)

QVQLVQSGAEVKKPGASVKVSCKASGYTFDYMNVWRQAPGQGLEWMGVISTFSGNTNFNQKFKGRVTMTTDTSTST  
AYMELRSLRSDDTAVYYCARSDYYGPYFDYWQGTTTLTVSSGGGGSGGGSGGGSDIVMTQTPLSLSVTPGPASIS  
SCRSSQTIVHSNGNTYLEWYLQKPGQSPQQLLIYKVSNRFSGVPRFSGSGSGTDFTLKISRVEAEDVGVYYCFQGS  
HVPFTFGGGTKVEIKRT (SEQ ID NO:245)

#### Humanized C3 single chain X4 (linker is IgG1 and IgG2 modified hinge region)

(DNA)

cagggttcagctggtgcagctctggagctgaggtgaagaagcctggggcctcagtggaaggtctcctgcaaggttctggt  
 tacacctttaccgactacgccatgaactgggtgcgacaggccctggacaagggttgagtggtgagtgatcagc  
 accttcagcggtaacacaaaacttcaaccagaagttcaagggcagagtcaccatgaccacagacacatccacgagcaca  
 gcctacatggagctgaggagcctgagatctgacgacacggcctgtattactgtgcgagaagcgactactacggccca  
 tacttcgactactggggccagggcaccaccctgaccgtgtccagcgataaaaaccataactaaaccgccccaaaccggcg  
 ccggaactgctgggtggtcctggtaccggtactggtgggtccgactattaaacctccgaaacctccgaaacctgctccg  
 aacctgctgggtggtccggatattgtgatgaccagactccactctctctgtccgtcaccctggacagccggcctcc  
 atctcctgcaggtctagtgcagaccattgtccatagtaatggaaacacctatgtggagtggtacctgcagaagccaggc  
 cagtcctccacagctcctgatctataaggtttccaaccgggttctctggagtgccagataggttcagtggtcagcggttca  
 gggacagatttcacactgaaaatcagccgggtggaggtgaggtggtgggttttattactgcttccaaggttagccac  
 gtgcctttcaccttcggcgagggaaccaaggtggagatcaaacgaact (SEQ ID NO:248)

(amino acids)

QVQLVQSGAEVVKPGASVKVSKASGYTFDYMNVWRQAPQGLEWMGVISTFSGNTNENQKFKGRVTMTTDTSTST  
 AYMELRSLRSDDTAVYYCARSDYYGPYFDYWGGTTLTVSSDKTHTKPPKPAPELLGGPGTGTGGPTIKPPKPPKPA  
 PNLLGGPDIVMTQTPLSLSVTPGQPASISCRSSQTIVHSNGNTYLEWYLQKPGQSPQLLIYKVSNRFSQVPDRFSGS  
 GSGTDFTLKISRVEAEDVGVYYCFQGSHPVFTFGGGTKVEIKRT (SEQ ID NO:249)

#### Humanized C8 single chain GS3 (linker is [Gly<sub>4</sub>Ser<sub>1</sub>]<sub>3</sub>)

(DNA)

gaggtgcagctggtggagctctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagtggtatgccatgagctgggtccgccagggtccagggaagggtggtgagtggtctcaaccattagt  
 agtggcggaacctacatatactacctgactcagtggaagggccgattcaccatctccagagacaacgccaagaactca  
 ctgtatctgcaaatgaacagcctgagagccgaggacacggcctgtattactgtgcgagactggcgcgcgataactat  
 tatgaatattggggcaaagggaaccacgggtcacctctcctccggcggtggcggtatccggcggtggcggtatccggcggt  
 ggcggtatccgacatcgtgatgaccagctctccagactccctggctgtgtctctggcgagagggccaccatcaactgc  
 agggccagcaagagtggttagcaccagcggtacagctacatgcaactggtaaccagcagaaaccaggacagcctcctaag  
 ctgctcatttacctggtgtctaacctggaatccgggggtccctgaccgattcagtggtcagcggtctgggacagatttc  
 actctcaccatcagcagcctgcaggtgaagatgtggcagtttattactgtcaacacattcgggaactgaccaggagt  
 gaattcggcgagggaaccaaggtggagatcaaacgaact (SEQ ID NO:250)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSGYAMSWVRQAPGKLEWVSTISSGGTYIYYPDSVKGRFTISRDNAAKN  
 SLYLQMNLSRAEDTAVYYCARLGGDNYYEYWGKGTITVTVSSGGGGSGGGSGGGGSDIVMTQSPDSLAVSLGERATI  
 NCRASKSVSTSGYSYMHWYQQKPGQPPKLLIYLVSNLESGVPDRFSGSGSGTDFTLTISSLQAEDVAVYYCQHIREL  
 TRSEFGGGTKVEIKRT (SEQ ID NO:251)

#### pSECTag2 E6 scFV-FC

(DNA)

atggagacagacacactcctgctatgggtactgctgctctgggttccaggttccactggtgacgcggcccagccggcc  
 gaggtgcagctggtggagctctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagtaggtatggcatgagctgggtccgccagggtccagggaagggtggtgagtggtctcaaccattagt  
 ggcgaggacactacatatactaccagactcagtggaagggccgattcaccatctccagagacaacgccaagaacacc  
 ctgtatctgcaaatgaacagcctgagagccgaggacacggctgtgtattactgtaccagagataactatggccgcaac  
 tatgattatggcatggattattggggccagggcaccctggtgaccgtgagcagcgcggtggcggtatccggcggtggc  
 ggatccggcggtggcggtatccgaaattgtgttgacacagctctccagccacctgtctttgtctccaggggaaagagcc  
 acctcacctgcagcgccaccagcagtggttagctacatccactggtaccaacagaggcctggccagagccccaggctc  
 ctcatctatagcacctccaacctggccagcgccatccagccaggttcagtggtcagtggtggtctgggagcgactacact  
 ctaccatcagcagcctagagcctgaagattttgagtttattactgtcagcagcgtagcagctccccctttcaccttt  
 ggagcgccaccaaagtggaaattaaagagcccaaatcttgtgacaaaactcacacatgccaccgtgccagcaccct  
 gaactcctggggggacgctcagtccttctctccccccaaaacccaaggacacctcatgatctcccgaccctgag  
 gtcacatgctggtggtggagctgagccacgaagacctgaggtcaagttcaactggtacgtggacggcggtggaggtg  
 cataatgccaaagacaaagccgcgggaggagcagtaaacagcagtaaccgtgtggtcagcgtcctcacctcctgcac  
 caggactggtgaatggcaaggagtacaagtgaaggtctccaacaaagccctccagcccccatcgagaaaaccatc  
 tccaaagccaaagggcagccccgagaaccacaggtgtacacctgcccccatcccgaggagagatgaccaagaaccag  
 gtcagcctgacctgctggtcaaaggcttctatccagcgacatcgccgtggagtggtgagagcaatggcgagccggag



aacaactacaagaccacgcctcccggtgctggactccgacggctccttcttctctacagcaagctcaccggtggacaag  
agcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggtctctgcacaaccactacacgcagaagagc  
ctctccctgtctccgggtaaatgataa (SEQ ID NO:256)

(amino acids)

METDTLLLWVLLLWVPGSTGDAAQPAEVQLVESGGGLVLPKPGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTI  
SGGGTYIYYPDSVKGRFTISRDNKNTLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLVTVSSGGGGSG  
GGSGGGGSEIVLTQSPATLSLSPGERATLTCSATSSVSYIHWHYQQRPGQSPRLLIYSTSNLASGIPARFSGSGSGS  
DYTLTISSLEPEDFAVYYCQQRSSSPFTFGSGTKVEIKEPKSCDKTHTCPPCPAPELLGGPSVFLFPPKPKDTLMIS  
RTPEVTCVVDVSHEDPEVKFNWYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAP  
IEKTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTTPVLDSDGSFFLYS  
KLTVDKSRWQQGNVSCSVMHEALHNHYTQKSLSLSPGK\*\* (SEQ ID NO:257)

#### Human IgG1 Fc sequence:

(DNA)

gagcccaaatcttgtgacaaaactcacacatgcccacgtgcccagcacctgaactcctggggggaccgtcagttctt  
ctcttccccccaaaacccaaggacacccctcatgatctcccggaacctgaggtcacatgcgtggtggtggacgtgagc  
cacgaagacctgaggtcaagttcaactggtacgtggacggcggtggaggtgcataatgccaagacaaagccgcgggag  
gagcagtacaacagcacgtaccgtgtggtcagcgtcctcacgtcctgcaccaggactggctgaatggcaaggagtag  
aagtgaaggtctccaacaaagccctcccagccccatcgagaaaaccatctccaaagccaaagggcagccccgagaa  
ccacaggtgtacacctgcccccatcccgggaggagatgaccaagaaccaggtcagcctgacctgacctgggtcaaaggc  
ttctatcccagcgacatcgccgtggagtgggagagcaatgggcagccggagagaacaactacaagaccacgcctcccgtg  
ctggactccgacggctccttcttctctacagcaagctcaccggtggacaagagcaggtggcagcaggggaacgtcttc  
tcatgctccgtgatgcatgaggtctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgataa  
(SEQ ID NO:272)

(amino acids)

EPKSCDKTHTCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVKFNWYVDGVEVHNAKTKPR  
EEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLV  
KGFYPSDIAVEWESNGQPENNYKTTTPVLDSDGSFFLYSKLTVDKSRWQQGNVSCSVMHEALHNHYTQKSLSLSPG  
K\*\* (SEQ ID NO:273)

#### Human IgG1 CH2-CH3 domain sequence:

(DNA)

ccgtgcccagcacctgaactcctggggggaccgtcagttcttcttcttccccccaaaacccaaggacacccctcatgatc  
tcccggaacctgaggtcacatgcgtggtggtggacgtgagccacgaagacctgaggtcaagttcaactggtacgtg  
gacggcggtggaggtgcataatgccaagacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtc  
ctcacgtcctgcaccaggactggctgaatggcaaggagtacaagtgaaggtctccaacaaagccctcccagcccc  
atcgagaaaaccatctccaaagccaaagggcagccccgagaaccacaggtgtacacctgcccccatcccgggaggag  
atgaccaagaaccaggtcagcctgacctgacctgggtcaaaggcttctatcccagcgacatcgccgtggagtgggagagc  
aatgggcagccggagagaacaactacaagaccacgcctcccgtgctggactccgacggctccttcttctctacagcaag  
ctcacgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggtctctgcacaaccac  
tacacgcagaagagcctctccctgtctccgggtaaatgataa (SEQ ID NO:274)

(amino acids)

PCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVKFNWYVDGVEVHNAKTKPREEQYNSTYRVVS  
VLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEW  
ESNGQPENNYKTTTPVLDSDGSFFLYSKLTVDKSRWQQGNVSCSVMHEALHNHYTQKSLSLSPGK\*\* (SEQ ID  
NO:275)

#### Human IgG1 CH3 domain sequence:

(DNA)

gggcagccccgagaaccacaggtgtacacctgcccccatcccgggaggagatgaccaagaaccaggtcagcctgacc  
tgacctgggtcaaaggcttctatcccagcgacatcgccgtggagtgggagagcaatgggcagccggagagaacaactacaag  
accacgcctcccgtgctggactccgacggctccttcttctctacagcaagctcaccggtggacaagagcaggtggcag

caggggaacgtcttctcatgctccgtgatgcatgaggctctgcacaaccactacacgcagaagagcctctccctgtct  
ccgggtaaatgataa (SEQ ID NO:276)

(amino acids)

GQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTTPVLDSDGSFFLYSKLTVDKSRW  
QQGNVFCFSVMHEALHNHYTQKSLSLSPGK\*\* (SEQ ID NO:277)

#### **CAR-T E6 CD8/CD8/CD28/CD3z sequence:**

N-CD8ls-huMNE6scFv-CD8ecd fragment- CD8 transmembrane- CD28- CD3zeta-C  
(DNA)

atggccctgcccgtgaccgcttttctgctgctccccctggcgctgctgctgcacgcgccagggccagaggtccagctgggt  
gagagtggcgggtgggctgggttaagcctggcgggtccctgcggtgagctgcgcgcgagtggtttactttcagccga  
tatgggatgagttgggtgcggaagctcccggaagaggtggaatgggtctcaacaatctccgggggggacacttac  
atctattaccccgactcagtcaggggagatttaccatttcacgagacaacgctaagaataccctgtatttgcagatg  
aattctctgagagcagaggacacagctgtttactattgtaccgcgacaactatggcaggaactacgactacggtatg  
gactattggggacaagggacattgggtacagtgcagctggcgcgggggcagcggaggaggaggcagcgggtgggggg  
ggcagcgagatagtgtcacgcagtcacccgcgactctcagctctctcacctggggaacgagctaccctgacgtgctct  
gctacctctcagtgctcatatattcactgggtatcagcaacggccccgggcagtcacctagattgctcatttatagtagc  
tctaactctggcctcaggtatccctgcacgattttctggatctgggttcaggttctgattacacctcactatctctagc  
ctggagcctgaagactttgcccgtttattactgccagcagaggtctagctccccattcacctttgggagtgaggaccaag  
gttgaaattaaaacgacaacccccggccccagaccaccaacgccagccccaccatcgccagccaacccctgtctctg  
agaccagaagcctgtaggcctgccgcgggtggagctgtgcacacaagaggactggatttcgctgtgatatctacatt  
tgggccccgctcgaggcacatgtggagtgtcctcctctccctgggtgattaccctgtactgcagaagcaagcggctct  
cggctcctgcattctgattacatgaacatgaccccaagaagaccaggccccaccaggaaacattaccagccctacgct  
ccgccacgcgacttcgctgctacccgtcccgcttaagttctcccgatcagccgacgcgcctgcttacaagcagggc  
cagaaccaactgtacaacgagctgaatctcggtagacgggaagagtaacgagctgttgacaaaacggagagggccgcgac  
ccagaaatgggcggcaagcctcgaggaaaaacccccaggagggactgtacaatgagttgcagaaagataagatggca  
gaagcttatagcgagatcggaatgaaggggaaaggagacgagggaaaggacacgagcgcctttatcagggcctgtcc  
acagcaacaaaagatacgtatgacgcctcccatatgcaggcacttccaccacgggtgataa  
(SEQ ID NO:297)

(amino acids)

MALPVTALLLLPLALLLHAARPEVQLVESGGGLVKGPSLRSLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGT  
YIYYPDSVKGRFTISRDNKNTLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLVTVSSGGGGSGGGGSG  
GGGSEIVLTQSPATLSLSPGERATLTCSATSSVSYIHQYQRPQSPRLLIYSTSNLASGIPARFSGSGSGSDYTLT  
ISSLEPEDFAVYYCQQRSSSPFTFGSGTKVEIKTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFAC  
DIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSRSADAP  
AYKQGGNQLYNEINLGRREEYDVLDRKRRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRRKGHDG  
LYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:298)

#### **CAR-T E6 CD8/CD8/4-1BB/CD3z sequence:**

N-CD8ls-huMNE6scFv-CD8ecd fragment- CD8 transmembrane- 4-1BB- CD3zeta-C  
(DNA)

atggccctgcccgtgaccgcttttctgctgctccccctggcgctgctgctgcacgcgccagggccagaggtccagctgggt  
gagagtggcgggtgggctgggttaagcctggcgggtccctgcggtgagctgcgcgcgagtggtttactttcagccga  
tatgggatgagttgggtgcggaagctcccggaagaggtggaatgggtctcaacaatctccgggggggacacttac  
atctattaccccgactcagtcaggggagatttaccatttcacgagacaacgctaagaataccctgtatttgcagatg  
aattctctgagagcagaggacacagctgtttactattgtaccgcgacaactatggcaggaactacgactacggtatg  
gactattggggacaagggacattgggtacagtgcagctggcgcgggggcagcggaggaggaggcagcgggtgggggg  
ggcagcgagatagtgtcacgcagtcacccgcgactctcagctctctcacctggggaacgagctaccctgacgtgctct  
gctacctctcagtgctcatatattcactgggtatcagcaacggccccgggcagtcacctagattgctcatttatagtagc  
tctaactctggcctcaggtatccctgcacgattttctggatctgggttcaggttctgattacacctcactatctctagc  
ctggagcctgaagactttgcccgtttattactgccagcagaggtctagctccccattcacctttgggagtgaggaccaag  
gttgaaattaaaacgacaacccccggccccagaccaccaacgccagccccaccatcgccagccaacccctgtctctg  
agaccagaagcctgtaggcctgccgcgggtggagctgtgcacacaagaggactggatttcgctgtgatatctacatt

tgggccccgctcgagggcacatgtggagtgctcctcctcctcctggtgattaccctgtactgcaaaaggggcccgc  
 aaactcctttacatttttaagcagccttttatgaggccagtagacagcactcaagaggaagacgggtgctcatgccgc  
 tttcctgaggaggaggaaggaggggtgcgaactgcgcgttaagttctcccgatcagccgacgcgcctgcttacaagcag  
 ggccagaaccaactgtacaacgagctgaatctcggtagacgggaagagtagcagctggttgacaaacggagagggccgc  
 gaccagaaatgggcggaagcctcgagggaaaaacccccaggagggactgtacaatgagttgcagaaagataagatg  
 gcagaagcttatagcgagatcggaatgaagggggaaaggagacgagggaaaggacacgacggcctttatcagggcctg  
 tccacagcaacaaaagatacgtatgacgcctccatattgcaggcacttccaccacggtgataa  
 (SEQ ID NO:300)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKGPGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGT  
 YIYYPDSVKGRFTISRDNKNTLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLVTVSSGGGSGGGGSG  
 GGGSEIVLTQSPATLSLSPGERATLTCSATSSVSYIHQYQQRPGQSPRLLIYSTSNLASGIPARFSGSGSGSDYTLT  
 ISSLEPEDFAVYYCQQRSSSPFTFGSGTKVEIKTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFAC  
 DIYIWAPLAGTCGVLLLSLVITLYCKRGRKKLLYIFKQPFMRPVQTTQEEDGCSCRFPEEEEGGCELRVKFSRSADA  
 PAYKQGQNQLYNELNLGRREEYDVLDRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRGKRGHD  
 GLYQGLSTATKDTYDALHMQALPPR\*\*  
 (SEQ ID NO:301)

#### **CAR-T E6 CD8/CD8/CD28/4-1BB/CD3z sequence:**

N-CD8ls-huMNE6scFv-CD8ecd fragment- CD8 transmembrane- CD28- 4-1BB- CD3zeta-C  
 (DNA)

atggccctgcccgtgaccgctttgctgctccccctggcgctgctgctgcaagcgcgcagggccagaggtccagctgggt  
 gagagtggcggtgggctggttaagcctggcggtcctcctgcggtgagctgcgccgcgagtggttactttcagccga  
 tatgggatgagttgggtgcggaagctcccggaagaggtggaatgggtctcaacaatctccggggggggcacttac  
 atctattaccccgactcagtcaggggagatttaccatttcacgagacaaacgctaagaataccctgtatttgcagatg  
 aattctctgagagcagaggacacagctgtttactattgtaccgcgcacaaactatggcaggaactacgactacggtatg  
 gactattggggacaagggacattggttacagtgagcagtgccggcggggggcagcggaggaggaggcagcgggtgggggg  
 ggacgcgagatagtgctcacgcagtcacccgcgactctcagtcctcaccctggggaacgagctaccctgacgtgctct  
 gctacctcctcagtgctcatatattcactggtatcagcaacggcccgggcagtcacctagatttgctcatttatagtagc  
 tctaactctggcctcaggtatccctgcacgattttctggatctgggttcaggttctgattacacctcactatctctagc  
 ctggagcctgaagacttttgcgtttattactgccagcagaggtctagctccccattcacccttgggagtgggaccaag  
 gttgaaattaaaacgacaacccccggccccccagaccaccaacgccagccccaccatcgccagccaacccctgtctctg  
 agaccagaagcctgtaggcctgcccgcgggtggagctgtgcacacaagaggactggatttcgcctgtgatattctacatt  
 tgggccccgctcgagggcacatgtggagtgctcctcctcctcctggtgattaccctgtactgcagaagcaagcggctc  
 cggctcctgcattctgattacatgaacatgaccccaagaagaccaggccccaccaggaaacattaccagccctacgct  
 ccgcccgcgacttctgctgctaccgggtccaaaagggggcgcaaaaaactcctttacatttttaagcagccttttatg  
 agggcagtagacgactcaagaggaagacgggtgctcatgccgctttcctgaggaggaggaaggagggtgcgaactg  
 cgcgttaagttctcccgatcagccgacgcgcctgcttacaagcagggccagaaccaactgtacaacgagctgaatctc  
 ggtagacgggaagagtagcagctggttgacaaacggagagggccgcgacccagaaatgggcggaagcctcgagggaaa  
 aacccccaggaggagtagtacaatgagttgcagaaagataagatggcagaagcttatagcgagatcggaatgaagggg  
 gaaaggagacgagggaaaggacacgacggcctttatcagggcctgtccacagcaacaaaagatacgtatgacgcctc  
 catatgcaggcacttccaccacggtgataa  
 (SEQ ID NO:303)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKGPGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGT  
 YIYYPDSVKGRFTISRDNKNTLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLVTVSSGGGSGGGGSG  
 GGGSEIVLTQSPATLSLSPGERATLTCSATSSVSYIHQYQQRPGQSPRLLIYSTSNLASGIPARFSGSGSGSDYTLT  
 ISSLEPEDFAVYYCQQRSSSPFTFGSGTKVEIKTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFAC  
 DIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYQPYAPPRDFAAYRSKRGRKKLLYIF  
 KQPFMRPVQTTQEEDGCSCRFPEEEEGGCELRVKFSRSADAPAYKQGQNQLYNELNLGRREEYDVLDRRGRDPEMG  
 GKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRGKRGHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID  
 NO:304)

**CAR-T C2 CD8/CD8/CD28/4-1BB/CD3z sequence:**

N-CD81s-huMNC2scFv-CD8ecd fragment- CD8 transmembrane- CD28- 4-1BB- CD3zeta-C  
(DNA)

atggccttgccagtgacggccctgctgctgccattggctcttctgttgacgctgccaggcctgaagtgcagctcgta  
gagagtggcggggactgggtgaagcccggtggaagcctcagactcagttgcccgcctcaggtttcactttttcaggt  
tacgccatgtcctgggtaagacaggcaccggggaaaggactcgagtgggtgtctactatcagctcaggaggcacttat  
atatattatcctgactctgtaaaaggccgattttacgattttctcgcgacaatgcaaagaactccctctacctccaaatg  
aacagtcttagggcagaagacactgctgtatactattgtgacgcctcgggcgacaaactactacgagtactttgac  
gtgtgggggaaagggactaccgtgacagtttcaagcggaggaggtggctcaggtggaggcggtcaggggggggagga  
agtgatattgtgctcacacaatccccagcctccctggctgtgtctcccggccaacgcgtacaattacatgtcggggc  
tccaaaagcgtgagcaccagcggctacagctacatgcactggtatcaacagaaaccaggacaaccccccaaactgttg  
atttatctcgttcaacttgagtcggcgctgctgcccgttttcagggagtgggagcggcacagattttacgctg  
actatcaaccccgtagaagcaaacgatacagcgaattattattgtcaacattcccgggaactccccctttacgttcggc  
gggggcacaaaggctcgaatttaagagaaccacgacaaccccgccccagaccaccaacgcagccccaccatcgcc  
agccaacccctgtctctgagaccagaagcctgtaggcctgcccgggtggagctgtgcacacaagaggactggatttc  
gcctgtgatattacatttggccccgctcgaggcacatgtggagtgtcctcctctcctcgttgattacctgtac  
tgcagaagcaagcgggtctcggctcctgcattctgattacatgaacatgaccccaagaagaccaggccccaccaggaaa  
cattaccagccctacgctccgcccagcgacttctgctgctaccgggtccaaaagggcgcaaaaaactcctttacatt  
tttaagcagccttttatgaggccagtacagacgactcaagaggaagacgggtgctcatgccgctttcctgaggaggag  
gaaggagggtgcgaactgcgcttaagttctcccgatcagccgacgcgctgcttacaagcaggggccagaaccaactg  
tacaacgagctgaatctcggtagacgggaagagtacgacgtgttggaacaacggagagggccgcgacccagaaatgggc  
ggcaagcctcgaggaaaaacccccaggaggactgtacaatgagttgcagaaagataagatggcagaagcttatagc  
gagatcggaatgaaggggaaaggagacgagggaaaggacacgacggcctttatcagggcctgtccacagcaacaaaa  
gatacgtatgacgcctccatattgcaggcacttccaccacgggtgataa (SEQ ID NO:306)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGTYIYPDSVKGRFTISRDNAKNS  
LYLQMNSLRAEDTAVYYCARLGGDNYYEYFDVWGKGTITVTVSSGGGGSGGGSGGGGSDIVLTQSPASLAVSPGQRAT  
ITCRASKSVSTSGYSYMHWYQKPGQPPKLLIYLASNLESGVPAFSGSGSGTDFTLTINPVEANDTANYYCQHSREL  
PFTFGGGTKVEIKRTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLLSL  
VITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYQPYAPPRDFAAYRSKRGRKLLYIFKQPFMRPVQTTQEEDGCSCR  
FPEEEEGGCELRVKFSRSADAPAYKQGQNQLYNELNLGRREEYDVLDKRRGRDPEMGGKPRRKNPQEGLYNELQKDKM  
AEAYSEIGMKGERRRGKGHDLGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:307)

**Humanized E6 scFV sequence in CAR:**

(DNA)

gaggtccagctgggtgagagtggcggtgggctgggttaagcctggcggtcctcgcggtgagctgcgcgcgagtgga  
tttactttcagccgatatgggatgagttgggtgcggcaagctcccgggaagaggctggaatgggtctcaacaatctcc  
ggggggggcacttacatctattaccccgactcagtcgaaggggagatttaccatttcacgagacaacgctaagaatacc  
ctgtatttgagatgaattctctgagagcagaggacacagctgttactattgtaccgcgacaactatggcaggaac  
tacgactacggtatggactattggggacaaggacatttggttacagtgagcagtgggcgggggggcagcggaggagga  
ggcagcgggtggggggggcagcagatagtgctcacgcagtcacccgcgactctcagctctctcacctggggaaacgagct  
acctgacgtgctctgctacctcctcagtgctcatatattcactggtatcagcaacggccccgggcagtcacctagattg  
ctcatttatagtagctctaattctggcctcaggtatccctgcagattttctggatctgggttcaggttctgattacacc  
ctcactatctctagcctggagcctgaagactttgcggtttattactgccagcagaggtctagctccccattcaccttt  
gggagtgggaccaaggttgaaattaaa (SEQ ID NO:341)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISSGGTYIYPDSVKGRFTISRDNAKN  
TLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLTVTVSSGGGGSGGGSGGGGSEIVLTQSPATLSLSPGE  
RATLTCSATSSVSIYHWYQRPQGSPRLLIYSTNLASGIPARFSGSGSGSDYTLTISSELPEDFAVYYCQQRSSSP  
FTFGSGTKVEIK (SEQ ID NO:342)

**CD8 leader sequence:**

(DNA)

atggccctgcccgtgaccgctttgctgctccccctggcgctgctgctgacgcgcgcaggcca (SEQ ID NO:343)

(amino acids)  
MALPVTALLLPLALLLHAARP (SEQ ID NO:344)

**CD8 hinge domain sequence:**

(DNA)  
acgacaacccccggccccagaccaccaacgccagccccaccatcgccagccaacccctgtctctgagaccagaagcc  
tgtaggcctgccgcgggtggagctgtgcacacaagaggactggatttcgcctgtgat (SEQ ID NO:345)

(amino acids)  
TTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACD (SEQ ID NO:346)

**CD4 hinge domain sequence:**

(DNA)  
tcgggacaggtcctgctggaatccaacatcaaggttctgcccacatgggtccacccccgggtgcagcca (SEQ ID  
NO:347)

(amino acids)  
SGQVLLESNIKVLPTWSTPVQP (SEQ ID NO:348)

**CD28 hinge domain sequence:**

(DNA)  
aaacacctttgtccaagtcacctatttcccgaccttctaagccc (SEQ ID NO:349)

(amino acids)  
KHLCPSPFLPGPSKP (SEQ ID NO:350)

**CD3 zeta transmembrane domain sequence:**

(DNA)  
ctctgctacctgctggatggaatcctcttcatctatgggtgtcattctcactgccttggttctg (SEQ ID  
NO:361)

(amino acids)  
LCYLLDGILFIYGVILTALFL (SEQ ID NO:362)

**CD8 transmembrane domain sequence:**

(DNA)  
atctacatttgggccccgctcgcaggcacatgtggagtgtcctcctctccctgggtgattaccctgtactgc (SEQ  
ID NO:363)

(amino acids)  
IYIWAPLAGTCGVLLLSLVITLYC (SEQ ID NO:364)

**CD4 transmembrane domain sequence:**

(DNA)  
atggccctgattgtgctggggggcgctcgccggcctcctgcttttcattgggctaggaatcttcttc (SEQ ID  
NO:365)

(amino acids)  
MALIVLGGVAGLLLLFIGLGIFF (SEQ ID NO:366)

**CD28 transmembrane domain sequence:**

(DNA)  
ttttgggtgctgggtgggtgggtggagtcctggcttgctatagcttgctagtaacagtggcctttattatcttg  
ggtg (SEQ ID NO:367)

(amino acids)

FWVLVVVGVLACYSLLVTVAFIIFWV (SEQ ID NO:368)

**4-1BB transmembrane domain sequence:**

(DNA)

atcatctccttctttcttgcgctgacgtcgactgcgttgctcttctctgctgttcttctcagcgtccggtttctctgt  
tggt (SEQ ID NO:369)

(amino acids)

IISFFLALTSTALLFLLFLLTLRFSVV (SEQ ID NO:370)

**OX40 transmembrane domain sequence:**

(DNA)

gttgccgccatcctgggcctgggcctggtgctggggctgctgggccccctggccatcctgctggccctgtacctgctc  
(SEQ ID NO:371)

(amino acids)

VAAILGLGLVLGLLGPLAILLALYLL (SEQ ID NO:372)

**CD3 zeta domain sequence:**

(DNA)

cgcgttaagttctcccgatcagccgacgcgcctgcttacaagcagggccagaaccaactgtacaacgagctgaatctc  
ggtagacgggaagagtacgacgtgttgacaaacggagagggcgcgacccagaaatgggcggcaagcctcgcaggaaa  
aacccccaggaggactgtacaatgagttgcagaaagataagatggcagaagcttatagcgagatcggaatgaagggg  
gaaaggagacgagggaaaggacacgacggcctttatcagggcctgtccacagcaacaaaagatacgtatgacgcctc  
catatgcaggcacttccaccacgg (SEQ ID NO:373)

(amino acids)

RVKF SRSADAPAYKQGQNQLYNELNLGRREEYDVLDKRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMK  
ERRRGKGHDGLYQGLSTATKDTYDALHMQALPPR (SEQ ID NO:374)

**CD3 zeta domain variant sequence:**

(DNA)

agagtgaagttcagcaggagcgcagacgccccgcgtaccagcagggccagaaccagctctataacgagctcaatct  
aggacgaagagaggagtacgatgttttgacaaagagacgtggccgggacccctgagatggggggaaagccgagaagga  
agaaccctcaggaaggcctgtacaatgaactgcagaaagataagatggcggaggcctacagtgagattgggatgaaa  
ggcgagcgccggaggggcaaggggcacgatggcctttaccaggggtctcagtacagccaccaaggacacctacgacgc  
ctttcacatgcaggccctgccccctcgc (SEQ ID NO:375)

(amino acids)

RVKF SRSADAPAYQQGQNQLYNELNLGRREEYDVLDKRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMK  
GERRRGKGHDGLYQGLSTATKDTYDALHMQALPPR (SEQ ID NO:376)

**CD28 domain sequence:**

(DNA)

agaagcaagcgggtctcggtcctgcattctgattacatgaacatgaccccaagaagaccagggccccaccaggaaacat  
taccagccctacgtctcgccacgcgacttcgctgctaccgggtcc (SEQ ID NO:377)

(amino acids)

RSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRS (SEQ ID NO:378)

**4-1BB domain sequence:**

(DNA)

aaaagggggccgcaaaaaactcctttacatttttaagcagccttttatgaggccagtacagacgactcaagaggaagac  
gggtgctcatgccgctttctgaggaggaggaaggagggtgcgaactg (SEQ ID NO:379)

(amino acids)

KRGRKKLLYIFKQPFMRPVQTTQEEDGCSCRFPEEEEGGCEL (SEQ ID NO:380)

**HumanizeE6 scFV (VH-VL) sequence:**

(DNA)

gaggtgcagctggtggagtctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagtaggtatggcatgagctgggtccgccaggctccagggaagaggctggagtgggtctcaaccattagt  
 ggcgaggacacatactatactacccagactcagtgaagggccgattcaccatctccagagacaacgccaaagacacc  
 ctgtatctgcaaatgaacagcctgagagccgaggacacggctgtgtattactgtaccagagataactatggccgcaac  
 tatgattatggcatggattattggggccagggcaccctggtgaccgtgagcagcgggcggtggcggtatccggcggtggc  
 ggatccggcggtggcggtatccgaaattgtgttgacacagctctccagccaccctgtctttgtctccaggggaaagagcc  
 accctcacctgcagcgccaccagcagtggttagctacatccactggtaccaacagaggcctggccagagccccaggctc  
 ctcatctatagcacctccaacctggccagcggtatccagccagggttcagtggcagtggtctgggagcgactacact  
 ctaccatcagcagcctagagcctgaagattttgcagtttattactgtcagcagcgtagcagctccccctttcaccttt  
 ggcagcggcaccaaagtggaaattaaa (SEQ ID NO:391)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGTYYYPDSVKGRFTISRDN  
 AKNTLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLTVTVSSGGGGSGGGSGGGGSEIVLTQSPATLSLSPGE  
 RATLTCSATSSVSYIHWHYQQRPGQSPRLLIYSTNLASGIPARFSGSGSGSDYTLTISSLEPEDFAVYYCQQRSSSP  
 FTFGSGTKVEIK (SEQ ID NO:392)

**HumanizeE6 scFV (VL-VH) sequence:**

(DNA)

gaaattgtgttgacacagctctccagccaccctgtctttgtctccaggggaaagagccaccctcacctgcagcgccacc  
 agcagtggttagctacatccactggtaccaacagaggcctggccagagccccaggctcctcatctatagcacctccaac  
 ctggccagcggtatccagccagggttcagtggcagtggtctgggagcgactacactctcaccatcagcagcctagag  
 cctgaagattttgcagtttattactgtcagcagcgtagcagctccccctttcacctttggcagcggtacccaaagtggaa  
 attaaaggcggtggcggtatccggcggtggcggtatccggcggtggcggtatccgaggtgcagctggtggagtctggggga  
 ggctggtcaagcctgggggggtccctgagactctcctgtgcagcctctggattcaccttcagtaggtatggcatgagc  
 tgggtccgccaggctccagggaagaggctggagtgggtctcaaccattagtggcgaggacacacatactaccca  
 gactcagtggaagggccgattcaccatctccagagacaacgccaaagacaccctgtatctgcaaatgaacagcctgaga  
 gccgaggacacggctgtgtattactgtaccagagataactatggccgcaactatgattatggcatggattattggggc  
 cagggcaccctggtgaccgtgagcagc (SEQ ID NO:393)

(amino acids)

EIVLTQSPATLSLSPGERATLTCSATSSVSYIHWHYQQRPGQSPRLLIYSTNLASGIPARFSGSGSGSDYTLTISSL  
 EPEDFAVYYCQQRSSSPFTFGSGTKVEIKGGGGSGGGSGGGGSEVQLVESGGGLVKPGGSLRLSCAASGFTFSRYG  
 MSWVRQAPGKRLEWVSTISGGGTYYYPDSVKGRFTISRDNKNTLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMD  
 YWGQGLTVTVSS (SEQ ID NO:394)

**HumanizeC2 scFV (VH-VL) sequence:**

(DNA)

gaggtgcagctggtggagtctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagtggtatgccatgagctgggtccgccaggctccagggaaggggctggagtgggtctcaaccattagt  
 agtggcggaacctacatactatactaccccgactcagtgaagggccgattcaccatctccagagacaacgccaaagactca  
 ctgtatctgcaaatgaacagcctgagagccgaggacacggcctgtattactgtgcgagacttgggggggataattac  
 tacgaataacttcgatgtctggggcaaagggaccacggtcaccgtctcctccggcggtggcggtatccggcggtggcgga  
 tccggcggtggcggtatccgacattgtgctgaccagctctccagcctccttggcctgtctccaggacagagggccacc  
 atcacctgcagagccagtaagagtgtcagtagcagcggtatcctacatgcactggtatcagcagaaaccaggacaa  
 cctcctaaactcctgatttacctggcatccaactctggagagcggggtccagccagggtcagcggcagtggtctggg  
 accgatttcaccctcacaattaatcctgtggaagctaatgatactgcaaattattactgtcagcagtagggagctg  
 cctttcacattcggcgaggggaccaaggtggagatcaaacgaact (SEQ ID NO:395)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSGYAMSWVRQAPGKLEWVSTISSGGTYYYPDSVKGRFTISRDN  
 AKNTLYLQMNSLRAEDTAVYYCARLGGDNYEYFDVWGKTTVTVSSGGGGSGGGSGGGGSDIVLTQSPASLAVSPGQR

ATITCRASKSVSTSGYSYMHWYQQKPGQPPKLLIYLASNLESGVPARFSGSGSGTDFTLTINPVEANDTANYYCQHS  
RELPTFTGGGTKVEIKRT (SEQ ID NO:396)

**HumanizeE6 scFV (VL-VH) sequence:**

(DNA)

gacattgtgctgacccagtcctccagcctccttggccgtgtctccaggacagagggccaccatcacctgcagagccagt  
aagagtgtcagtagccagcggatactcctacatgcactggatcagcagaaaccaggacaacctcctaaactcctgatt  
tacctggcatccaatctggagagcgggggtccagccaggttcagcggcagtggggtctgggaccgatttcacctcaca  
attaatcctgtggaagctaatactgatactgcaaattattactgtcagcacagtagggagctgcctttcacattcggcgga  
gggaccaaggtggagatcaaacgaactggcgggtggcggatccggcgggtggcggatccggcgggtggcggatccgaggtg  
cagctgggtggagctctgggggagggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctggattcacc  
ttcagtggtatgccatgagctgggtccgcccaggtccagggaaggggctggagtgggtctcaaccattagtagtggc  
ggaacctacatatactaccccgactcagtggaagggccgattcaccatctccagagacaacgccaagaactcactgtat  
ctgcaaatagaacagcctgagagccgaggacacggccgtgtattactgtgcgagacttgggggggataattactacgaa  
tacttcgatgtctggggcaaagggaccacgggtcaccgtctcctcc (SEQ ID NO:397)

(amino acids)

DIVLTQSPASLAVSPGQRATITCRASKSVSTSGYSYMHWYQQKPGQPPKLLIYLASNLESGVPARFSGSGSGTDFTL  
TINPVEANDTANYYCQHSRELPTFTGGGTKVEIKRTGGGGSGGGGSGGGGSEVQLVESGGGLVKPGGSLRLSCAASG  
FTFGYAMSWVRQAPGKLEWVSTISSGGTYIYYPDSVKGRFTISRDNKNSLYLQMNSLRAEDTAVYYCARLGGDN  
YYEYFDVWGKTTTVTVSS (SEQ ID NO:398)

**G<sub>4</sub>S<sub>1</sub> linker sequence:**

(DNA)

ggcgggtggcggatcc (SEQ ID NO:399)

(amino acids)

GGGGS (SEQ ID NO:400)

**[G<sub>4</sub>S<sub>1</sub>]<sub>x3</sub> linker sequence:**

(DNA)

ggcgggtggcggatccggcgggtggcggatccggcgggtggcggatcc (SEQ ID NO:401)

(amino acids)

GGGSGGGSGGGGS (SEQ ID NO:402)

**8 aa GS linker sequence:**

(DNA)

ggcgggttcggcgggtggatccgga (SEQ ID NO:403)

(amino acids)

GGSGGGSG (SEQ ID NO:404)

**12 aa GS linker sequence:**

(DNA)

ggcgggttcggcgggtggatccggcgggtggcggatccgga (SEQ ID NO:405)

(amino acids)

GGSGGGSGGGSG (SEQ ID NO:406)

**13 aa GS linker sequence:**

(DNA)

ggcgggtggatccggcgggtggcggatccggcgggtggatcc (SEQ ID NO:407)

(amino acids)

GGSGGGSGGGGS (SEQ ID NO:408)



**22 aa GS linker sequence:**

(DNA)

ggcgggtggaagcggcggtggcggatccggcagcggcggaagcggcggtggcggatccggcggtgga (SEQ ID NO:409)

(amino acids)

GGSGGGSGSGSGSGSGSGG (SEQ ID NO:4110)

**24 aa GS linker sequence:**

(DNA)

ggcggttccggcggtggatccggcggtggcggatccggaggcgggtccggcggtggatccggcggtggcggatccgga (SEQ ID NO:411)

(amino acids)

GGSGGGSGGGSGGGSGGGSGG (SEQ ID NO:412)

**Mouse C3 Heavy chain variable region sequence:**

(DNA)

caggtccagctgcagcagctctgggcctgagctggtgaggcctgggggtctcagtgaaagatttctgcaagggttccggc  
tacagattcactgattatgctatgaactgggtgaagcagagtcatgcaaagagtcctagagtggttgagttattagtc  
actttctctggttaatacaaaacttcaaccagaagtttaagggcaaggccacaatgactgtagacaaatcctccagcaca  
gcctatatggaacttgccagattgacatctgaggattctgccatgtattactgtgcaagatcggattactacggccca  
tactttgactactggggccaaggcaccactctcacagtctcctca (SEQ ID NO:413)

(amino acids)

QVQLQQSGPELVRPGVSVKISCKGSGYRFTDYAMNWKQSHAKSLEWIGVISTFSGNTNFNQKFKGKATMTVDKSSST  
AYMELARLTSEDSAMYYCARSDYYGPYFDYWQGTTTLTVSS (SEQ ID NO:414)**Mouse C3 heavy chain variable framework region 1 (FWR1) sequence:**

(DNA)

caggtccagctgcagcagctctgggcctgagctggtgaggcctgggggtctcagtgaaagatttctgcaagggttccgg  
ctacagattcact (SEQ ID NO:415)

(amino acids)

QVQLQQSGPELVRPGVSVKISCKGSGYRFT (SEQ ID NO:416)

**Mouse C3 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

gattatgctatgaac (SEQ ID NO:417)

(amino acids)

DYAMN (SEQ ID NO:418)

**Mouse C3 heavy chain variable framework region 2 (FWR2) sequence:**

(DNA)

tgggtgaagcagagtcatgcaaagagtcctagagtggttgga (SEQ ID NO:419)

(amino acids)

WVKQSHAKSLEWIG (SEQ ID NO:420)

**Mouse C3 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

gttattagttactttctctggttaatacaaaacttcaaccagaagtttaagggc (SEQ ID NO:421)

(amino acids)

VISTFSGNTNFNQKFKG (SEQ ID NO:422)

**Mouse C3 heavy chain variable framework region 3 (FWR3) acid sequence:**

(DNA)

aaggccacaatgactgtagacaaatcctccagcacagcctatatggaacttgccagattgacatctgaggattctgc  
catgtattactgtgcaaga (SEQ ID NO:423)

(amino acids)

KATMTVDKSSSTAYMELARLTSEDSAMYYCAR (SEQ ID NO:424)

**Mouse C3 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

tcggattactacggcccatactttgactac (SEQ ID NO:425)

(amino acids)

SDYYGPYFDY (SEQ ID NO:426)

**Humanized C3 heavy chain variable region sequence from IGHV1-18\*04:**

(DNA)

caggttcagctggtgcagctctggagctgaggtgaagaagcctggggcctcagtggaaggtctcctgcaaggcttctgg  
ttacacctttaccgactacgccatgaactgggtgacagggccctggacaagggcttgagtggatgggagtgatca  
gcaccttcagcggtaacacaaaacttcaaccagaagttcaagggcagagtcacatgaccacagacacatccacgagc  
acagcctacatggagctgaggagcctgagatctgacgacacggcgtgtattactgtgcgagaagcgactactacgg  
cccatacttcgactactggggccagggcaccacctgaccgtgtccagc (SEQ ID NO:439)

(amino acids)

QVQLVQSGAEVKKPGASVKVSCKASGYTFTDYAMNWVRQAPGQGLEWMGVISTFSGNTNFNQKFKGRVTMTTDTSTS  
TAYMELRSLRSDDTAVYYCARSDYYGPYFDYWGQGTTLTVSS (SEQ ID NO:440)

**Humanized C3 heavy chain variable framework region 1 (FWR1) acid sequence:**

(DNA)

caggttcagctggtgcagctctggagctgaggtgaagaagcctggggcctcagtggaaggtctcctgcaaggcttctgg  
ttacacctttacc (SEQ ID NO:441)

(amino acids)

QVQLVQSGAEVKKPGASVKVSCKASGYTFT (SEQ ID NO:442)

**Humanized C3 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

gactacgccatgaac (SEQ ID NO:443)

(amino acids)

DYAMN (SEQ ID NO:444)

**Humanized C3 heavy chain variable framework region 2 (FWR2) acid sequence:**

(DNA)

tgggtgacagggccctggacaagggcttgagtggatggga (SEQ ID NO:445)

(amino acids)

WVRQAPGQGLEWMG (SEQ ID NO:446)

**Humanized C3 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

gtgatcagcaccttcagcggtaacacaaaacttcaaccagaagttcaagggc (SEQ ID NO:447)

(amino acids)  
VISTFSGNTNFNQKFKG (SEQ ID NO:448)

**Humanized C3 heavy chain variable framework region 3 (FWR3) acid sequence:**

(DNA)  
agagtcaccatgaccacagacacatccacgagcacagcctacatggagctgaggagcctgagatctgacgacacggcc  
gtgtattactgtgcgaga (SEQ ID NO:449)

(amino acids)  
RVTMTTDTSTSTAYMELRSLRSDDTAVYYCAR (SEQ ID NO:450)

**Humanized C3 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)  
agcgactactacggccatacttcgactac (SEQ ID NO:451)

(amino acids)  
SDYYGPYFDY (SEQ ID NO:452)

**Humanized C3 IgG1 heavy chain sequence**

(DNA)  
caggttcagctggtgcagctctggagctgaggtgaagaagcctggggcctcagtggaaggtctcctgcaaggcttctggt  
tacacctttaccgactacgccatgaactgggtgcgacaggccctggacaagggttgagtggatgggagtgatcagc  
accttcagcggtaacacaaaacttcaaccagaagttcaagggcagagtcaccatgaccacagacacatccacgagcaca  
gcctacatggagctgaggagcctgagatctgacgacacggccgtgtattactgtgcgagaagcgactactacggccca  
tacttcgactactggggccagggcaccaccctgaccgtgtccagcgcctagcaccaaggggcccatcggtcttccccctg  
gcacctcctccaagagcacctctgggggcacagcggccctgggtgctcaaggactacttccccgaaccgggtg  
acggtgtcgtggaactcaggcgcctgaccagcggcgtgcacaccttcccggtgtcctacagtcctcaggactctac  
tccctcagcagcgtggtgacagtgcctccagcagcttgggcacccagacctacatctgcaacgtgaatcacaagccc  
agcaacaccaaggtggacaagaaagttagcccaaatcttgtaaaaaactcacacatgccaccgtgcccagcacct  
gaactcctgggggaccgtcagctcttctcttcccccaaaacccaaggacacctcatgatctcccggaacctgag  
gtcacatgcgtgggtgggtgacgtgagccacgaagacctgaggtcaagttcaactggtacgtggacggcgtggaggtg  
cataatgccaaagacaaagccgcgggaggagcagtacaacagcacgtaccgtgtgggcagcgtcctcaccgtcctgcac  
caggactggctgaatggcaaggagtacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaaccatc  
tccaaagccaaagggcagccccgagaaccacaggtgtacacctgcccccatcccgggaggagatgaccaagaaccag  
gtcagcctgacctgctggtcaaaggcttctatcccagcgacatcgccgtggagtgaggagcaatgggcagccggag  
aacaactacaagaccacgctcccggtgctggactccgacggctccttcttctctacagcaagctcaccgtggacaag  
agcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggctctgcacaaccactacacgcagaagagc  
ctctccctgtctccgggtaaatgataa (SEQ ID NO: 453)

(amino acids)  
QVQLVQSGAEVKKPGASVKVSKASGYTFTDYAMNWVRQAPGQGLEWMGVISTFSGNTNFNQKFKGRVTMTTDTSTS  
TAYMELRSLRSDDTAVYYCARSDYYGPYFDYWGQGTTLTVSSASTKGPSVFPLAPSSKSTSGGTAALGLVKDYFPE  
PVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVTVTPSSSLGTQTYICNVNHKPSNTKVDKKVEPKSCDKTHTCPPC  
PAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVKFNWYVDGVEVHNAKTKPREEQYNSTYRVVSVL  
TVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWES  
NGQPENNYKTTPVLDSDGSFFLYSKLTVDKSRWQQGNVFCSCVMHEALHNHYTQKSLSLSPGK\*\* (SEQ ID  
NO:454)

**Humanized C3 IgG2 heavy chain sequence**

(DNA)  
caggttcagctggtgcagctctggagctgaggtgaagaagcctggggcctcagtggaaggtctcctgcaaggcttctggt  
tacacctttaccgactacgccatgaactgggtgcgacaggccctggacaagggttgagtggatgggagtgatcagc  
accttcagcggtaacacaaaacttcaaccagaagttcaagggcagagtcaccatgaccacagacacatccacgagcaca  
gcctacatggagctgaggagcctgagatctgacgacacggccgtgtattactgtgcgagaagcgactactacggccca  
tacttcgactactggggccagggcaccaccctgaccgtgtccagcgcctccaccaaggggcccatcggtcttccccctg

gcgccctgctccaggagcacctccgagagcacagccgcccctgggctgctgggtcaaggactacttccccgaaccgggtg  
acgggtgctcgtggaactcaggcgctctgaccagcggcggtgcacaccttcccagctgtcctacagtcctcaggactctac  
tcctcagcagcgtgggtgaccgtgccctccagcaacttcggcaccagacctacacctgcaacgtagatcacaagccc  
agcaacaccaaggtggacaagacagttgagcgcgaatggtgtgctgagtgcccaccgtgccagcaccacctgtggca  
ggaccgtcagtccttcttccccccaaaacccaaggacacctcatgatctcccggaccctgaggtcacgtgctg  
gtgggtggacgtgagccacgaagacccccgaggtccagttcaactggtacgtggacggcggtggaggtgcataatgccaa  
gacaagccacgggaggagcagttcaacagcacgttccgtgtgggtcagcgtcctcaccgttgtgcaccaggactggctg  
aacggcaaggagtacaagtgaaggtctccaacaaaggcctcccagcccccatcgagaaaaccatctccaaaacccaaa  
gggcagccccgagaaccacaggtgtacacctgcccccatcccgggaggagatgaccaagaaccaggtcagcctgacc  
tgctggtcaaaggcttctaccccagcgacatcgccgtggagtgaggaggaatgggcagccggagaacaactacaag  
accacacctcccatgctggactccgacggctccttcttctctacagcaagctcaccgtggacaagagcaggtggcag  
caggggaacgtcttctcatgctccgtgatgcatgaggtctctgcacaaccactacacgcagaagagcctctccctgtct  
ccgggtaaatagtaa (SEQ ID NO:455)

(amino acids)

QVQLVQSGAEVKKPGASVKVSKASGYTFTDYAMNWVRQAPGQGLEWMGVISTFSGNTNFNQKFKGRVTMTTDTSTS  
TAYMELRSLRSDDTAVYYCARSDYYGPYFDYWGQGTTLTVSSASTKGPSVFPLAPCSRSTSESTAALGCLVKDYFPE  
PVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVTVPSNFGTQTYTCNVDHKPSNTKVDKTVKCCVECPPCPAP  
PVAGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVQFNWYVDGVEVHNAKTKPREEQFNSTFRVSVLTVVH  
QDWLNGKEYKCKVSNKGLPAPIEKTISKTKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQP  
ENNYKTTTPMLDSDGSFFLYSKLTVDKSRWQQGNVFCSCVMHEALHNHYTQKSLSLSPGK\*\* (SEQ ID NO:456)

**Mouse C3 Light Chain variable region sequence:**

(DNA)

gatgttttgatgacccaaactccactctccctgctgctcagtccttggagatcaagcctccatctcttgcagatctagt  
cagaccattgtacatagtaatggaaacacctatttagaatggtacctgcagaaaccaggccagtcctccaaagctcctg  
atctacaaagtttccaaccgattttctgggtcccagacaggttcagtggcagtgagtcagggacagatttcacactc  
aagatcaacagagtgagggtgaggtctgggagtttattactgctttcaaggttcacatgttccattcacgttcggc  
tcggggacaaagttggaataaaaa (SEQ ID NO:458)

(amino acids)

DVLMTQTPLSLPVSLGDQASISCRSSQTIVHSNGNTYLEWYLQKPGQSPKLLIYKVSNRFSGVPRDRFSGSGSGTDFTL  
KINRVEAEDLGVIYCFQGSHVPFTFGSGTKLEIK (SEQ ID NO:459)

**Mouse C3 light chain variable framework region 1 (FWR1) sequence :**

(DNA)

gatgttttgatgacccaaactccactctccctgctgctcagtccttggagatcaagcctccatctcttgc (SEQ ID  
NO:460)

(amino acids)

DVLMTQTPLSLPVSLGDQASISC (SEQ ID NO:461)

**Mouse C3 light chain variable complementarity determining regions 1 (CDR1)  
sequence :**

(DNA)

agatctagtcagaccattgtacatagtaatggaaacacctatttagaa (SEQ ID NO:462)

(amino acids)

RSSQTIVHSNGNTYLE (SEQ ID NO:463)

**Mouse C3 light chain variable framework region 2 (FWR2) sequence:**

(DNA)

tggtacctgcagaaaccaggccagtcctccaaagctcctgatctac (SEQ ID NO:464)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:465)

**Mouse C3 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

aaagtttccaaccgattttct (SEQ ID NO:466)

(amino acids)

KVSNRFS (SEQ ID NO:467)

**Mouse C3 light chain variable framework region 3 (FWR3) sequence:**

(DNA)

gggggtcccagacaggttcagtggtcagtggtcagggacagatttcacactcaagatcaacagagtgagggtgagga  
tctgggagtttattactgc (SEQ ID NO:468)

(amino acids)

GVPDRFSGSGSGTDFTLKINRVEAEDLGVIYC (SEQ ID NO:469)

**Mouse C3 light chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

tttcaaggttcacatgttccattcacg (SEQ ID NO:470)

(amino acids)

FQGSHPFT (SEQ ID NO:471)

**Humanized C3 light chain variable region sequence from IGKV2-29\*03:**

(DNA)

gatattgtgatgaccagactccactctctctgtccgtcaccctggacagccggcctccatctcctgcaggtctagt  
cagaccattgtccatagtaatggaacacctatttggagtggtacctgcagaagccaggccagtcctccacagctcctg  
atctataaggtttccaaccggttctctgtgagtgccagataggttcagtggtcagcggtcagggacagatttcacactg  
aaaatcagccgggtggaggtgaggtgaggtggtgggtttattactgttccaaggtagccacgtgcctttcaccttcggc  
ggagggaccaaggtggagatcaaacgaact (SEQ ID NO:486)

(amino acids)

DIVMTQTPLSLSVTPGQPASISCRSSQTIVHSNGNTYLEWYLQKPGQSPQLLIYKVSNRFSGVPDRFSGSGSGTDFT  
LKISRVEAEDVGVYYCFQGSHPFTFGGGTKVEIKRT (SEQ ID NO:487)**Humanized C3 light chain variable framework region 1 (FWR1) acid sequence:**

(DNA)

gatattgtgatgaccagactccactctctctgtccgtcaccctggacagccggcctccatctcctgc (SEQ ID  
NO:488)

(amino acids)

DIVMTQTPLSLSVTPGQPASIS (SEQ ID NO:489)

**Humanized C3 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

ggtctagtcagaccattgtccatagtaatggaacacctatttggag (SEQ ID NO:490)

(amino acids)

RSSQTIVHSNGNTYLE (SEQ ID NO:491)

**Humanized C3 light chain variable framework region 2 (FWR2) acid sequence:**

(DNA)

tggtacctgcagaagccaggccagtcctccacagctcctgatctat (SEQ ID NO:492)

(amino acids)

WYLQKPGQSPQLLIY (SEQ ID NO:493)

**Humanized C3 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

aaggtttccaaccggttctct (SEQ ID NO:494)

(amino acids)

KVSNRFS (SEQ ID NO:495)

**Humanized C3 light chain variable framework region 3 (FWR3) acid sequence:**

(DNA)

ggagtgccagataggttcagtggcagcgggtcagggacagatttcacactgaaaatcagccgggtggaggctgaggatgttggggtttattactgc (SEQ ID NO:496)

(amino acids)

GVPDRFSGSGSGTDFTLKISRVEAEDVGVYYC (SEQ ID NO:497)

**Humanized C3 light chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

ttccaaggtagccacgtgcctttcacc (SEQ ID NO:498)

(amino acids)

FQGSHPFT (SEQ ID NO:499)

**Humanized C3 lambda light chain sequence**

(DNA)

gatattgtgatgaccagactccactctctctgtccgtcaccctggacagcggcctccatctcctgcaggtctagt  
cagaccattgtccatagtaatggaaacacctatttggagtggtagctgcagaagccaggccagtcctccacagctcctg  
atctataaggtttccaaccggttctctggagtgccagataggttcagtggcagcgggtcagggacagatttcacactg  
aaaatcagccgggtggaggctgaggatgttgggggtttattactgttccaaggtagccacgtgcctttcaccttcggc  
ggagggaaccaaggtggagatcaaacgaactggcagcccaaggctgccccctcggtcactctgttcccgccctcctct  
gaggagcttcaagccaacaaggccacactgggtgtgtctcataagtgaacttctacccgggagccgtgacagtgccctgg  
aaggcagatagcagccccgtcaaggcgggagtggagaccaccacacctccaaacaaagcaacaacaagtaacgcggcc  
agcagctatctgagcctgacgctgagcagtggaagtccacagaagctacagctgccagggtcacgcataaagggagc  
accgtggagaagacagtggtccctacagaatgttcatagtaa (SEQ ID NO:500)

(amino acids)

DIVMTQTPLSLSVTPGPASISCRSSQTIVHSNGNTYLEWYLQKPGQSPQLLIYKVSNNRFSVDPDRFSGSGSGTDFT  
LKISRVEAEDVGVYYCFQGSHPFTFGGGTKVEIKRTGQPKAAPSVTLFPPSSEELQANKATLVCLISDFYPGAVTV  
AWKADSSPVKAGVETTPSKQSNKYYAASSYLSLTPEQWKSHRYSYSCQVTHEGSTVEKTVAPTECS\*\* (SEQ ID  
NO:501)

**Humanized C3 Kappa light chain**

(DNA)

gatattgtgatgaccagactccactctctctgtccgtcaccctggacagcggcctccatctcctgcaggtctagt  
cagaccattgtccatagtaatggaaacacctatttggagtggtagctgcagaagccaggccagtcctccacagctcctg  
atctataaggtttccaaccggttctctggagtgccagataggttcagtggcagcgggtcagggacagatttcacactg  
aaaatcagccgggtggaggctgaggatgttgggggtttattactgttccaaggtagccacgtgcctttcaccttcggc  
ggagggaaccaaggtggagatcaaacgaactacgggtggctgcaccatctgttctatcttcccgccatctgatgagcag  
ttgaaatctggaactgcctctgttgtgtgctgctgaataacttctatcccagagaggccaaagtacagtggaaggtg  
gataacgccttccaatcgggtaactcccaggagagtgtcacagagcaggacagcaaggacagcacctacagcctcagc  
agcacctgacgctgagcaagcagactacgagaaacacaaagtctacgctgcgaagtaccccatcagggcctgagc  
tcgcccgtcacaaagagcttcaacaggggagagtgttagtaa (SEQ ID NO:502)

(amino acids)

DIVMTQTPLSLSVTPGQPASISCRSSQTIVHSNGNTYLEWYLQKPGQSPQLLIYKVSNRFSGVDPDRFSGSGSGTDFT  
 LKISRVEAEDVGVYYCFQGSHVPFTFGGGTKVEIKRTTVAAPSVFIFPPSDEQLKSGTASVVCLLNNFYPREAKVQW  
 KVDNALQSGNSQESVTEQDSKSTYLSSTLTLSKADYEKHKVYACEVTHQGLSSPVTKSFNRGEC\*\* (SEQ ID  
 NO: 503)

**Mouse C8 heavy chain variable region sequence**

(DNA)

gaagtgatggctcgtggaaagcggcggtggtctggttaaagcgggggggatcccttaagctttcttgcgccgcacatccggg  
 ttcacgttctccggctatgccatgtcctgggtccgacagactcccgaaaagcgttggaatgggtggccactatctcc  
 tccggggggacgtacatctactaccccgacagtggtgaaaggaagatttacaatatctcgcgacaacgcaaaaaatacc  
 ttgtatcttcaaatagagtcctcgtcggtcagaggacactgccatgtactattgcgcccgcctgggcggcgacaattac  
 tatgagtat (SEQ ID NO:505)

(amino acids)

EVMVVESGGGLVKPGGSLKLSAASGFTFSGYAMSWVRQTPEKRLEWVATISSGGTYIYYPDSVKGRFTISRDNKNT  
 LYLQMSSLRSEDTAMYYCARLGDNYYEY (SEQ ID NO:506)

**Mouse C8 heavy chain variable complementarity determining region 1 (CDR1)  
 sequence:**

(DNA)

ggctatgccatgtcc (SEQ ID NO:507)

(amino acids)

GYAMS (SEQ ID NO:508)

**Mouse C8 heavy chain variable complementarity determining region 2 (CDR2)  
 sequence:**

(DNA)

actatctcctccggggggacgtacatctactaccccgacagtggtgaaagga (SEQ ID NO:509)

(amino acids)

TISSGGTYIYYPDSVKG (SEQ ID NO:510)

**Mouse C8 heavy chain variable complementarity determining region 3 (CDR3)  
 sequence:**

(DNA)

ctgggcggcgacaattactatgagtat (SEQ ID NO:511)

(amino acids)

LGDNYYEY (SEQ ID NO:512)

**Humanized C8 heavy chain variable region sequence from IGHV3-21\*04:**

(DNA)

gaggtgcagctggtggagctctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagtggctatgccatgagctgggtccgccaggtccaggggaaggggtggagtggtctcaaccattagt  
 agtggcggaacctacatatactacctgactcagtgaaagggcgattcaccatctccagagacaacgccaagaactca  
 ctgtatctgcaaatgaacagcctgagagccgaggacacggcgtgtattactgtgcgagactgggcggcgataactat  
 tatgaatattggggcaaagggaccacgggtcacctctcctcc (SEQ ID NO: 525)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSGYAMSWVRQAPGKLEWVSTISSGGTYIYYPDSVKGRFTISRDNKNT  
 SLYLQMNSLRAEDTAVYYCARLGDNYYEYWGKGTITVTVSS (SEQ ID NO: 526)

**Humanized C8 heavy chain variable framework region 1 (FWR1) sequence:**

(DNA)

gaggtgcagctggtggagctctgggggaggcctggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
 ttcaccttcagt (SEQ ID NO:527)

(amino acids)  
EVQLVESGGGLVKPGGSLRLSCAASGFTFS (SEQ ID NO:528)

**Humanized C8 heavy chain variable complementarity determining region 1 (CDR1) sequence:**

(DNA)  
ggctatgccatgagc (SEQ ID NO:529)

(amino acids)  
GYAMS (SEQ ID NO:530)

**Humanized C8 heavy chain variable framework region 2 (FWR2) sequence:**

(DNA)  
tgggtccgccaggctccaggaaggggctggagtgggtctca (SEQ ID NO:531)

(amino acids)  
WVRQAPGKGLEWVS (SEQ ID NO:532)

**Humanized C8 heavy chain variable complementarity determining region 2 (CDR2) sequence:**

(DNA)  
accattagtagtggcggaacctacatatactaccctgactcagtgaagggc (SEQ ID NO:533)

(amino acids)  
TISSGGTYIYPDSVKG (SEQ ID NO:534)

**Humanized C8 heavy chain variable framework region 3 (FWR3) sequence:**

(DNA)  
cgattcaccatctccagagacaacgccaaagaactcactgtatctgcaaatgaacagcctgagagccgaggacacggcc  
gtgtattactgtgcgaga (SEQ ID NO:535)

(amino acids)  
RFTISRDNAKNSLYLQMNSLRAEDTAVYYCAR (SEQ ID NO:536)

**Humanized C8 heavy chain variable complementarity determining region 3 (CDR3) sequence:**

(DNA)  
ctgggcggcgataactattatgaatat (SEQ ID NO:537)

(amino acids)  
LGGDNYEY (SEQ ID NO:538)

**Humanized C8 IgG1 heavy chain sequence**

(DNA)  
gaggtgcagctggtggagtctgggggaggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
ttcaccttcagtggctatgccatgagctgggtccgccaggctccaggaaggggctggagtgggtctcaaccattagt  
agtggcggaacctacatatactaccctgactcagtgaagggccgattcaccatctccagagacaacgccaaagaactca  
ctgtatctgcaaatgaacagcctgagagccgaggacacggccgtgtattactgtgcgagactgggcggcgataactat  
tatgaatatattggggcaaagggaaccacgggtcacctctcctccgctagcaccgaaggcccatcggtcttccccctggca  
ccctcctccaagagcacctctgggggcacagcggccctgggtgctggtcaaggactactccccgaaccgggtgacg  
gtgtcgtggaactcaggcgccctgaccagcggcggtgcacacctcccggtgtctactacagtcctcaggactctactcc  
ctcagcagcgtggtgacagtgccctccagcagcttgggcaccacagacctacatctgcaacgtgaatcacaagcccagc  
aacaccaaggtggacaagaaagttgagcccaaatcttgtgacaaaactcacacatgccacccgtgccagcacctgaa  
ctcctgggggggaccgtcagtccttctcttccccccaaaacccaaggacacctcatgatctcccggaaccttgaggtc  
acatgcgtggtggtggacgtgagccacgaagacctgaggtcaagttcaactggtacgtggacggcggtggaggtgcat  
aatgccaagacaaagccgcgaggagcagtagacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccag  
gactggctgaatggcaaggagtacaagtgaaggtctccaacaagccctccagcccccatcgagaaaaccatctcc



aaagccaaagggcagccccgagaaccacaggtgtacacctgcccccatcccgggaggagatgaccaagaaccaggtc  
agcctgacctgacctgggtcaaaggcttctatcccagcgacatcgccgtggagtgggagagcaatgggcagccggagaac  
aactacaagaccacgcctcccgtgctggactccgacggctccttcttctctacagcaagctcaccgtggacaagagc  
aggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggctctgcacaaccactacacgcagaagagcctc  
tcctgtctccgggtaaatgataa (SEQ ID NO:539)

(amino acids)

EVQLVESGGGLVPGGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGTYIYYPDSVKGRFTISRDNKNS  
LYLQMNSLR AEDTAVYYCARLGGDNYEYWGKGTITVTVSSASTKGPSVFPLAPSSKSTSGGTAALGCLVKDYFPEPVT  
VSWNSGALTSGVHTFPAVLQSSGLYSLSSVTVPSSSLGTQTYICNVNHKPSNTKVDKKVEPKSCDKTHTCPPCPAPE  
LLGGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVKFNWYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQ  
DWLNGKEYKCKVSNKALPAPIEKTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN  
NYKTTTPVLDSDGSFFLYSKLTVDKSRWQQGNVFCSCVMHEALHNHYTQKSLSLSPGK\*\* (SEQ ID NO:540)

#### Humanized C8 IgG2 heavy chain sequence

(DNA)

gaggtgcagctgggtggagtctgggggaggcctgggtcaagcctgggggggtccctgagactctcctgtgcagcctctgga  
ttcaccttcagtggctatgccatgagctgggtccgccaggtccagggaaggggtggagtgggtctcaaccattagt  
agtggcggaacctacataactacctgactcagtgaaggcgcatcaccatctccagagacaacgccagaactca  
ctgtatctgcaaatgaacagcctgagagccgaggacacggcctgtattactgtgcgagactgggcggcgataactat  
tatgaatattggggcaaagggaccacgggtcacctgtctcctccgctccaccaagggcccatcggtcttccccctggcg  
ccctgtccaggagcacctccgagagcacagccgccccgggtgctgctggtcaaggactacttccccgaaccgggtgacg  
gtgtcgtggaactcaggcgctctgaccagcggcgtgcacaccttcccagctgtcctacagtcctcaggactctactcc  
ctcagcagcgtgggtgaccgtgccctccagcaacttcgggcacccagacctacacctgcaacgtagatcacaagcccagc  
aacaccaaggtggacaagacagttgagcgcgaatgttggtgctgagtgcccacogtggccagcaccacctgtggcgagga  
ccgtcagctcttctcttcccccccaaaacccaaggacacctcatgatctcccggacccctgaggtcacgtgcgtgggtg  
gtggacgtgagccacgaagaccccgaggtccagttcaactggtaegtggagcggcgtggaggtgcataatgccaagaca  
aagccacgggaggagcagttcaacagcaggttcctgtggtcagcgtcctcaccgttggtgcaccaggactggctgaac  
ggcaaggagtacaagtgaaggtctccaacaaaggcctcccagcccccatcgagaaaacatctccaaaaccaaaggg  
cagccccgagaaccacaggtgtacacctgcccccatcccgggaggagatgaccaagaaccaggtcagcctgacctgc  
ctggtcaaaggcttctaccccagcgacatcgccgtggagtgggagagcaatgggcagccggagacaactacaagacc  
acacctccatgctggactccgacggctccttcttctctacagcaagctcaccgtggacaagagcaggtggcagcag  
gggaacgtcttctcatgctccgtgatgaggtctctgcacaaccactacacgcagaagagcctctccctgtctccg  
ggtaaatagtaa (SEQ ID NO:541)

(amino acids)

EVQLVESGGGLVPGGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGTYIYYPDSVKGRFTISRDNKNS  
LYLQMNSLR AEDTAVYYCARLGGDNYEYWGKGTITVTVSSASTKGPSVFPLAPCSRSTSESTAALGCLVKDYFPEPVT  
VSWNSGALTSGVHTFPAVLQSSGLYSLSSVTVPSNFGTQTYTCNVDHKPSNTKVDKTKVERKCCVECPPCPAPPVAG  
PSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVQFNWYVDGVEVHNAKTKPREEQFNSTFRVVSVLTVVHQDWLN  
GKEYKCKVSNKGLPAPIEKTISKTKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKT  
TPPMLDSDGSFFLYSKLTVDKSRWQQGNVFCSCVMHEALHNHYTQKSLSLSPGK\*\* (SEQ ID NO:542)

#### Mouse C8 light chain variable region sequence

(DNA)

gacatcgctcattacgcagacccctgccagctcttgccgtttctctggggccagagggccactatcagttacagggcgagt  
aagtctgtgagtaccagcggctatagttacatgcattggaaccagcagaaacggggacagccaccacgcctgcttatt  
tatctgggtgctaatcttgagtcgggggtgcccgccaggttcagcggcagcggctctgggaccgacttcacactcaac  
attcatccagtggaagaagaggacgtgctacatactactgtcaacacattcgggaactgaccaggagtgaa (SEQ  
ID NO:543)

(amino acids)

DIVITQTPASLAVSLGQRATISYRASKSVSTSGYSYMHWNQQKPGQPRLLIYLVSNLESGVPARFSGSGSGTDFTLN  
IHPVEEEDAATYYCQHIRELTRSE (SEQ ID NO:544)

#### Mouse C8 light chain variable complementarity determining region 1 (CDR1) sequence:

(DNA)

aggggcgagtaagtctgtgagtaccagcggctatagttacatgcat (SEQ ID NO:545)

(amino acids)

RASKSVSTSGYSYM (SEQ ID NO: 546)

**Mouse C8 light chain variable complementarity determining region 2 (CDR2) sequence:**

(DNA)

ctgggtgtctaatacttgagtcc (SEQ ID NO:547)

(amino acids)

LVSNLES (SEQ ID NO:548)

**Mouse C8 light chain variable complementarity determining region 3 (CDR3) sequence:**

(DNA)

caacacattcgggaactgaccaggagtga (SEQ ID NO:549)

(amino acids)

QHIRELTRSE (SEQ ID NO:550)

**Humanized C8 light chain variable region sequence from NCBI germline z00023:**

(DNA)

gacatcgatgaccagtcctccagactccctggctgtgtctctgggcgagagggccaccatcaactgcagggccagc  
aagagtgttagcaccagcggctacagctacatgcactggtagcagcagaaccaggacagcctcctaagctgctcatt  
tacctgggtgtctaacctggaatccggggctccctgacagattcagtggcagcgggtctgggacagatttcactctcacc  
atcagcagcctgcaggctgaagatgtggcagtttattactgtcaacacattcgggaactgaccaggagtgaattcggc  
ggagggaccaaggtggagatcaaacgaact (SEQ ID NO:565)

(amino acids)

DIVMTQSPDSLAVSLGERATINCRASKSVSTSGYSYMHYQQKPGQPPKLLIYLVSNLESGVPDRFSGSGSGTDFTL  
TISSLQAEDVAVYYCQHIRELTRSEFGGGTKVEIKRT (SEQ ID NO:566)

**Humanized C8 light chain variable framework region 1 (FWR1) sequence:**

(DNA)

gacatcgatgaccagtcctccagactccctggctgtgtctctgggcgagagggccaccatcaactgc (SEQ ID NO:567)

(amino acids)

DIVMTQSPDSLAVSLGERATINC (SEQ ID NO:568)

**Humanized C8 light chain variable complementarity determining region 1 (CDR1) sequence:**

(DNA)

agggccagcaagagtgttagcaccagcggctacagctacatg (SEQ ID NO:569)

(amino acids)

RASKSVSTSGYSYM (SEQ ID NO:570)

**Humanized C8 light chain variable framework region 2 (FWR2) sequence:**

(DNA)

cactggtagcagcagaaccaggacagcctcctaagctgctcatttac (SEQ ID NO:571)

(amino acids)

HWYQQKPGQPPKLLIY (SEQ ID NO:572)

**Humanized C8 light chain variable complementarity determining region 2 (CDR2) sequence:**

(DNA)

ctggtgtctaacctggaatcc (SEQ ID NO:573)

(amino acids)

LVSNLES (SEQ ID NO:574)

**Humanized C8 light chain variable framework region 3 (FWR3) sequence:**

(DNA)

ggggtccttgaccgattcagtgggcagcgggtctgggacagatttcactctcaccatcagcagcctgcaggctgaagatgtggcagtttattactgt (SEQ ID NO:575)

(amino acids)

GVPDRFSGSGSGTDFTLTISSLQAEDVAVYYC (SEQ ID NO:576)

**Humanized C8 light chain variable complementarity determining region 3 (CDR3) sequence:**

(DNA)

caacacattcggaactgaccaggagtga (SEQ ID NO:577)

(amino acids)

QHIRELTRSE (SEQ ID NO:578)

**Humanized C8 Lambda light chain sequence**

(DNA)

gacatcgatgacctcagactccctggctgtgtctctgggogagagggccaccatcaactgcagggccagcaagagtgttagcaccagcggctacagctacatgcactggtagcagcagaaaccaggacagcctcctaagctgctcatttacctgggtgtctaacctggaatccggggtccttgaccgattcagtgggcagcgggtctgggacagatttcactctcaccatcagcagcctgcaggctgaagatgtggcagtttattactgtcaacacattcgggaactgaccaggagtgaattcggcggagggaccaaggtggagatcaaacgaactggtagcccaaggctgccccctcggtcactctgttccccccctcctctgaggagcttcaagccaacaggccacactgggtgtgtctcataagtgacttctaccgggagccgtgacagtggcctggaagcagatagcagccccgtcaaggcgggagtgagaccaccacacctccaaacaaagcaacaacaagtacgcggccagcagctatctgagcctgagcctgagcagtggaagtccacagaagctacagctgccagggtcacgcatgaagggagcaccgtggagaagacagtgggccctacagaatgttcatagtaa (SEQ ID NO:579)

(amino acids)

DIVMTQSPDSLAVSLGERATINCRASKSVSTSGYSYMHWYQQKPGQPPLLIYLVSNLESGVPDRFSGSGSGTDFTLTISSLQAEDVAVYYCQHIRELTRSEFGGGTKVEIKRTGQPKAAPSVTLFPPSSEELQANKATLVCLISDFYPGAVTVAWKADSSPVKAGVETTPSKQSNNKYAASSYLSLTPEQWKSHRSYSCQVTHEGSTVEKTVAPTECS\*\* (SEQ ID NO:580)

**Humanized C8 Kappa light chain sequence**

(DNA)

gacatcgatgacctcagactccctggctgtgtctctgggogagagggccaccatcaactgcagggccagcaagagtgttagcaccagcggctacagctacatgcactggtagcagcagaaaccaggacagcctcctaagctgctcatttacctgggtgtctaacctggaatccggggtccttgaccgattcagtgggcagcgggtctgggacagatttcactctcaccatcagcagcctgcaggctgaagatgtggcagtttattactgtcaacacattcgggaactgaccaggagtgaattcggcggagggaccaaggtggagatcaaacgaactacgggtggctgcaccatctgtcttcattctccccccatctgatgagcagttgaaatctggaactgcctctgttgtgctgctgaataacttctatcccagagagggccaaagtacagtggaaaggtgataacgccccccaatcggttaactcccaggagagtggtcacagagcaggacagcaaggacagcacctacagcctcagcagcacctgacgtgagcaaacgagactacgagaaacacaaagtctacgcctgcgaagtcacccatcagggcctgagctcgccccgtcacaaagagcttcaacaggggagagtgtagtaa (SEQ ID NO:581)

(amino acids)

DIVMTQSPDSLAVSLGERATINCRASKSVSTSGYSYMHWYQQKPGQPPLLIYLVSNLESGVPDRFSGSGSGTDFTLTISSLQAEDVAVYYCQHIRELTRSEFGGGTKVEIKRTTVAAPSVFIFPPSDEQLKSGTASVVCLLNNFYPREAKVQWKV

DNALQSGNSQESVTEQDSKDYSLSTLTLSKADYEKHKVYACEVTHQGLSSPVTKSFNRGEC\*\* (SEQ ID NO:582)

#### CAR-T E6 CD8 sequence:

(DNA)

gaggtccagctggttgagagtggtggtgggctggttaagcctggcggctccctgcggtgagctgcgcgcgagtgga  
tttactttcagccgatatgggatgagttgggtgcggcaagctcccggaagaggtggaatgggtctcaacaatctcc  
ggggggggcacttacatctattaccccgactcagtcaggggagatttaccatttcacgagacaacgctaagaatacc  
ctgtatttgcagatgaattctctgagagcagaggacacagctgtttactattgtacccgcgacaactatggcaggaac  
tacgactacggtatggactattggggacaagggacattggttacagtgcagctggcggcgggggcagcggaggagga  
ggcagcgggtggggggggcagcagatagtgtcagcagtcacccgcgactctcagctctctcacctggggaacgagct  
accctgacgtgctctgctacctcctcagtgctcatatattcactggtatcagcaacggcccgggcagctcccttagattg  
ctcatttatagtagcttaattctggcctcaggtatccctgcacgattttctggatctgggtcaggttctgattacacc  
ctcactatctctagcctggagcctgaagactttgcccgtttattactgccagcagaggtctagctccccattcaccttt  
gggagtgaggaccaaggttgaaattaaaacgacaaccccgccccagaccaccaacgccagccccaccatcgccagc  
caaccctgtctctgagaccagaagcctgtaggcctgccgcgggtggagctgtgcacacaagaggactggatttcgcc  
tgtgatattctacatttggggcccgctcgcaggcacatgtggagtgtcctcctctccctggtgattaccctgtactgc  
tgataa (SEQ ID NO:584)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGTYYYPDSVKGRFTISRDN  
AKN TLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLTVTVSSGGGGSGGGSGGGGSEIVLTQSPATLSLSPGE  
RATLTCSATSSVSIYHWYQQRPGQSPRLLIYSTSNLASGIPARFSGSGSGSDYTLTISLSEPEDFAVYYCQQRSSP  
FTFGSGTKVEIKTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLSLVI  
TLYC\*\* (SEQ ID NO:585)

#### CAR-T C2 CD8 CD8 sequence:

N-CD81s-huMNC2scFv-CD8ecd fragment- CD8 transmembrane-C

(DNA)

gaagtgcagctcgtagagagtggtggggggactggtgaagcccggtggaagcctcagactcagttgcgcgcctcaggt  
ttcactttttcaggttacgccatgtcctgggtgaagacaggcaccggggaaaggactcgagtggtgtctactatcagc  
tcaggaggcacttatatatattatcctgactctgtaaaaggccgatttaccgatttctcgcgacaatgcaaagaactcc  
ctctacctccaaatgaacagtccttagggcagaagacactgctgtatactattgtgcacgcctcggcgggcacaactac  
tacgagtactttgacgtgtgggggaaaggactaccgtgacagtttcaagcggaggaggtgggtcaggtggaggcggg  
tcaggggggggaggaagtgatattgtgctcacacaatccccagcctccctggctgtgtctcccgcccaacgcgctaca  
attacatgtcgggcctccaaaagcgtgagcaccagcggctacagctacatgcactggtatcaacagaaaccaggacaa  
ccccccaaactgttgatttatctcgttcaaaacttgaggctcggcgctgctgcgcgcttttcaggagtgaggagcggc  
acagattttacgctgactatcaaccccgtagaagcaaacgatacagcgaattattattgtcaacattcccggaactc  
ccctttacgttcggcgggggcacaaggtcgaaattaagagaaccacgacaaccccgccccagaccaccaacgcca  
gccccaccatcgccagccaacccctgtctctgagaccagaagcctgtaggcctgccgcgggtggagctgtgcacaca  
agaggactggatttcgctgtgatattctacatttggggcccgctcgcaggcacatgtggagtgtcctcctctccctg  
tgattaccctgtactgtctgataa (SEQ ID NO:586)

(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTFSGYAMSWVRQAPGKLEWVSTISSGGTYYYPDSVKGRFTISRDN  
AKN SLYLQMNSLRAEDTAVYYCARLGGDNYEYFDVWGKGTTVTVSSGGGGSGGGSGGGGSDIVLTQSPASLAVSPGQR  
ATITCRASKSVSTSGYSYHWHYQKPGQPPKLLIYLASNLESGVPARFSGSGSGTDFTLTINPVEANDTANYCQHS  
RELPTFGGGTKVEIKRTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLL  
LSLVITLYC\*\* (SEQ ID NO:587)

#### CD8/4-1BB sequence

N- CD8 transmembrane- 4-1BB-C

(DNA)

acgacaaccccgggccccagaccaccaacgccagccccaccatcgccagccaacccctgtctctgagaccagaagcc  
tgtaggcctgccgcgggtggagctgtgcacacaagaggactggatttcgctgtgatattctacatttggggcccgctc  
gcaggcacatgtggagtgtcctcctctcctcctggtgattaccctgtactgcaaaagggggccgcaaaaaactcctttac

atttttaagcagccttttatgaggccagtacagacgactcaagaggaagaacgggtgctcatgccgctttcctgaggag  
gaggaaggagggtgcgaactgtgataa (SEQ ID NO:588)

(amino acids)

TTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLLLSLVITLYCKRGRKKLL  
YIFKQPFMRPVQTTQEEEDGCSRFPEEEEGGCEL\*\* (SEQ ID NO:589)

#### CD8/CD28 sequence

N- CD8 transmembrane- CD28-C

(DNA)

acgacaacccccggccccagaccaccaacgccagccccaccatcgccagccaacccctgtctctgagaccagaagcc  
tgtaggcctgccgcccgtggagctgtgcacacaagaggactggatttcgcctgtgatattctacatttgggccccgctc  
gcaggcacatgtggagtgtcctcctctccttgggtgattaccctgtactgcagaagcaagcgggtctcggctcctgcat  
tctgattacatgaacatgaccccaagaagaccaggccccaccaggaacattaccagccctacgctccgccacgcgac  
ttcgctgcctaccgggtcctgataa (SEQ ID NO: 590)

(amino acids)

TTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLLLSLVITLYCRSKRSRLL  
HSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRS\*\* (SEQ ID NO:591)

#### CD8/CD3z sequence:

N- CD8 transmembrane- CD3zeta-C

(DNA)

acgacaacccccggccccagaccaccaacgccagccccaccatcgccagccaacccctgtctctgagaccagaagcc  
tgtaggcctgccgcccgtggagctgtgcacacaagaggactggatttcgcctgtgatattctacatttgggccccgctc  
gcaggcacatgtggagtgtcctcctctccttgggtgattaccctgtactgcgcggttaagttctcccgatcagccgac  
gcgcctgtttacaagcagggccagaaccaactgtacaacgagctgaatctcggtagacgggaagagtagcagctgttg  
gacaaacggagaggccgcgacccagaaatggggcggaagcctcgcaggaaaaacccccaggagggactgtacaatgag  
ttgcagaaagataagatggcagaagcttatagcgagatcggaatgaagggggaaaggagacgagggaaaggacacgac  
ggcctttatcaggggcctgtccacagcaacaaaagatacgtatgacgcctccatattgcaggcacttccaccacgggtga  
taa (SEQ ID NO:592)

(amino acids)

TTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLLLSLVITLYCRVKFSRSA  
DAPAYKQGQNQLYNELNLGRREEYDVLDRKRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRGKG  
HDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:593)

#### CD8/CD28/CD3z sequence:

N- CD8 transmembrane- CD28- CD3zeta-C

(DNA)

acgacaacccccggccccagaccaccaacgccagccccaccatcgccagccaacccctgtctctgagaccagaagcc  
tgtaggcctgccgcccgtggagctgtgcacacaagaggactggatttcgcctgtgatattctacatttgggccccgctc  
gcaggcacatgtggagtgtcctcctctccttgggtgattaccctgtactgcagaagcaagcgggtctcggctcctgcat  
tctgattacatgaacatgaccccaagaagaccaggccccaccaggaacattaccagccctacgctccgccacgcgac  
ttcgctgcctaccgggtcccgcggttaagttctcccgatcagccgacgcgcgctgtttacaagcagggccagaaccaactg  
tacaacgagctgaatctcggtagacgggaagagtagcagctgttggaacaaacggagagggccgcgacccagaaatgggc  
ggcaagcctcgcaggaaaaacccccaggagggactgtacaatgagttgcagaaagataagatggcagaagcttatagc  
gagatcggaatgaagggggaaaggagacgagggaaaggacacgacggcctttatcagggcctgtccacagcaacaaaa  
gatacgtatgacgcctccatattgcaggcacttccaccacgggtgataa (SEQ ID NO:594)

(amino acids)

TTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLLLSLVITLYCRSKRSRLL  
HSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSRSADAPAYKQGQNQLYNELNLGRREEYDVLDRKRRGRDPE  
MGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRGKGHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ  
ID NO:595)

#### CD8/4-1BB/CD3z sequence:

N- CD8 transmembrane- 4-1BB- CD3zeta-C

(DNA)

acgacaacccccggccccccagaccaccaacgccagccccccaccatcgccagccaacccctgtctctgagaccagaagcc  
tgtaggcctgccgcccgtggagctgtgcacacaagaggactggatttcgctgtgatattctacatttgggccccgctc  
gcaggcacatgtggagtgtcctcctctcctcctgggtgattaccctgtactgcaaaaggggcccgaaaaaactcctttac  
atttttaagcagccttttatgaggccagtagacagcactcaagaggaagacgggtgctcatgccgctttcctgaggag  
gaggaaggaggggtgcgaactgcgcgttaagttctcccgatcagccgacgcgcctgcttacaagcagggccagaaccaa  
ctgtacaacgagctgaatctcggtagacgggaagagtagcagcgtggttgacaaacggagagggccgcgacccagaaatg  
ggcggaagcctcgcaggaaaaacccccaggagggactgtacaatgagttgcagaaagataagatggcagaagcttat  
agcgagatcggaatgaagggggaaaggagacgagggaaaggacagcagcgccctttatcagggcctgtccacagcaaca  
aaagatacgtatgacgcctccatattgcaggcacttccaccacgggtgataa (SEQ ID NO:596)

(amino acids)

TTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLLLSLVITLYCKRGRKKLL  
YIFKQPFMRPVQTTQEEDGCSCRFPEEEEGGCELRVKFSRSADAPAYKQGQNQLYNELNLGRREEYDVLDKRRGRDP  
EMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRGKGHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ  
ID NO:597)

#### CD8/CD28/4-1BB/CD3z sequence:

N- CD8 transmembrane- CD28- 4-1BB- CD3zeta-C

(DNA)

acgacaacccccggccccccagaccaccaacgccagccccccaccatcgccagccaacccctgtctctgagaccagaagcc  
tgtaggcctgccgcccgtggagctgtgcacacaagaggactggatttcgctgtgatattctacatttgggccccgctc  
gcaggcacatgtggagtgtcctcctctcctcctgggtgattaccctgtactgcagaagcaagcgggtctcggctcctgcac  
tctgattacatgaacatgaccccaagaagaccaggccccaccaggaaacattaccagccctacgctccgcccacgcgac  
tctcgtgcttaccgggtccaaaaggggcccgaaaaaactcctttacatttttaagcagccttttatgaggccagtagacag  
acgactcaagaggaagacgggtgctcatgccgctttcctgaggaggaggaaggaggggtgcgaactgcgcgttaagttc  
tcccgatcagccgacgcgcctgcttacaagcagggccagaaacactgtacaacgagctgaatctcggtagacgggaa  
gagtacgacgtggttgacaaacggagagggccgcgacccagaaatggcgcggaagcctcgcaggaaaaacccccaggag  
ggactgtacaatgagttgcagaaagataagatggcagaagcttatagcgagatcggaatgaagggggaaaggagacga  
gggaaaggacacgacggcctttatcagggcctgtccacagcaacaaaagatacgtatgacgcctccatattgcaggca  
cttccaccacgggtgataa (SEQ ID NO:598)

(amino acids)

TTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLLLSLVITLYCRSKRSRL  
HSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSKRGRKKLLYIFKQPFMRPVQTTQEEDGCSCRFPEEEEGGCELRV  
KFSRSADAPAYKQGQNQLYNELNLGRREEYDVLDKRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGE  
RRRGKGHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO: 599)

#### CAR-T C2 CD8/CD8/CD28/CD3z sequence:

N-CD8ls-huMNC2scFv-CD8ecd fragment- CD8 transmembrane- CD28- CD3zeta-C

(DNA)

atggccttgccagtgacggccctgctgctgccattggctcttctgttgcaagctgccaggcctgaagtgcagctcgta  
gagagtggcgggggactgggtgaagcccgggtggaagcctcagactcagttgcgcgcctcaggtttcactttttcaggt  
tacgccatgtcctgggtaagacaggcaccggggaaaggactcgagtgggtgtctactatcagctcaggaggcacttat  
atatattatcctgactctgtaaaaggccgatttacgatttctcgcgacaatgcaaagaactccctctacctccaaatg  
aacagtcttagggcagaagacactgctgtatactattgtgcacgcctcggcgggcgacaactactacgagtactttgac  
gtgtgggggaaagggactaccgtgacagtttcaagcggaggaggtggctcaggtggaggcggtcaggggggggagga  
agtgtatattgtgtcacacaatccccagcctccctggctgtgtctcccgccaacgcgctacaattacatgtcgggccc  
tccaaaagcgtgagcaccagcggctacagctacatgcactgggtatcaacagaaaccaggacaacccccccaaactgttg  
atattctcgtcttcaaacttgagtcggcgctgctgcgcgcttttcagggagtgaggagcggcacagattttacgctg  
actatcaaccccgtagaagcaaacgatacagcgaattattattgtcaacattcccggggaactccccctttacgttcggc  
gggggcacaaaggtcgaaattaagagaaccacgacaacccccggccccccagaccaccaacgccagccccccaccatcgcc  
agccaacccctgtctctgagaccagaagcctgtaggcctgccgcccgtggagctgtgcacacaagaggactggatttc  
gcctgtgatattctacatttgggccccgctcgcaggcacatgtggagtgtcctcctctcctcctgggtgattaccctgtac

tgcagaagcaagcgggtctcggtcctgcattctgattacatgaacatgaccccaagaagaccaggccccaccaggaaa  
cattaccagccctacgctccgccacgcgacttctgctgctaccgggtcccgcttaagttctcccgatcagccgacgcg  
cctgcttacaagcagggccagaaccaactgtacaacgagctgaatctcggtagacgggaagagtacgacgtgttgac  
aaacggagagggccgcgacccagaaatggcgccgaagcctcgaggaaaaacccccaggagggactgtacaatgagttg  
cagaaagataagatggcagaagcttatagcgagatcggaatgaagggggaaaggagacgagggaaaggacacgacggc  
ctttatcagggcctgtccacagcaacaaaagatacgtatgacgcctccatattgcaggcacttccaccacgggtgataa  
(SEQ ID NO:608)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKGPGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGT  
YIYPDSVKGRFTISRDNAKNSLYLQMNSLRAEDTAVYYCARLGGDNYEYFDVWGKGTITVTVSSGGGSGGGGSGG  
GGSDIVLTQSPASLAVSPGQRATITCRASKSVSTSGYSYMHWYQQKPGQPPKLLIYLASNLESGVPARFSGSGSGTD  
FTLTINPVEANDTANYYCQHSRELPTFTGGGKVEIKRTTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTR  
GLDFACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFS  
RSADAPAYKQGQNQLYNELNLGRREEYDVLDRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRR  
KGKHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:609)

#### **CAR-T C2 CD8/CD8/4-1BB/CD3z sequence #13:**

N-CD8ls-huMNC2scFv-CD8ecd fragment- CD8 transmembrane- 4-1BB- CD3zeta-C  
(DNA)

atggccttgccagtgcggccctgctgctgccattggctcttctgttgacgctgccaggcctgaagtgcagctcgta  
gagagtggcggggactggtgaagcccggtggaagcctcagactcagttgcgcgcctcagggttctacttttccaggt  
tacgccatgtcctgggtaagacaggcaccggggaaggactcgagtgggtgtctactatcagctcaggaggcacttat  
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(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKGPGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGT  
YIYPDSVKGRFTISRDNAKNSLYLQMNSLRAEDTAVYYCARLGGDNYEYFDVWGKGTITVTVSSGGGSGGGGSGG  
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RSADAPAYKQGQNQLYNELNLGRREEYDVLDRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERR  
RGKHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:611)

#### **MUC1 truncated extra cellular domain sequence**

(amino acids)

SNIKFRPGSVVVQLTLAFREGTINVHDTVETQFNQYKTEAASRY (SEQ ID NO:620)

#### **MUC1 truncated extra cellular domain sequence**

(amino acids)

SVVVQLTLAFREGTINVHDTVETQFNQYKTEAASRY (SEQ ID NO:621)

**MUC1 truncated extra cellular domain sequence**

(amino acids)

VQLTLAFREGTINVHVDVETQFNQY (SEQ ID NO:622)

**MUC1 truncated extra cellular domain sequence**

(amino acids)

SNIKFRPGSVVVQLTALAFREGTIN (SEQ ID NO:623)**Primers**

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**MMP 9**

(DNA)

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(amino acids)

MSLWQPLVLVLVLLGCCFAAPRQRQSTLVLPDGLRTNLTDRQLAEEYLYRYGYTRVAEMRGESKSLGPALLLLQKQ  
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APPTVCPTGPPTVHPSERPSTAGPTGPPSAGPTGPPTAGPSTATTVPLSPVDDACNVNIFDAIAEIGNQLYLFKDGKY  
WRFSEGRGSRPQGPFLIADKWPALPRKLDVFEERLSKKLFFFSGRQVWVYTGA SVLGPRRLDKLGLGADVAQVTGA  
LRSGRGKMLLFSGRRLWRF DVKAQMVDPMSASEVDRMFPGVPLDTHDVFQYREKAYFCQDRFYWRVSSRSELNQVDQ  
VGYVTYDILQCPEDDYKDDDDK\*\* (SEQ ID NO:643)

### MMP9 catalytic domain

(DNA)

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(amino acids)

MFQTFEGDLKWHHHNITYWIQNYSEDLPRVIDDAFARAFALWSAVTPLTFTRVYSRDADIVIQFGVAEHGDGYPFDD  
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### NFATc1 Promoter (NFATc1P)

(DNA)

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211

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### NFATc1P-MMP9cat

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**NFAT response element**

(DNA)

ggaggaaaaactgtttcatacagaaggcgt (SEQ ID NO:649)

**NFAT response element repeats**

(DNA)

ggaggaaaaactgtttcatacagaaggcgtggaggaaaaactgtttcatacagaaggcgtggaggaaaaactgtttca  
tacagaaggcgt (SEQ ID NO:650)**CMV minimal promoter**

(DNA)

aggtaggcgtgtacgggtgggaggtctatataagcagagctggtttagtgaaccgtcagatc (SEQ ID NO:651)

**NFATREmCMV-MMP9**

(DNA)

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**NFATREmCMV-MMP9cat**

(DNA)

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## C2 scFv

(DNA)

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(amino acids)

EVQLVESGGGLVKGPGSLRLSCAASGFTFSGYAMSWVRQAPGKLEWVSTISSGGTYIYYPDSVKGRFTISRDNAKN  
 SLYLQMNSLRAEDTAVYYCARLGGDNYEYFDVWGKGTITVTVSSGGGGSGGGSGGGSDIVLTQSPASLAVSPGQR  
 ATITCRASKSVSTSGYSYMHYQQKPGQPPKLLIYLASNLESGVPARFSGSGSGTDFTLTINPVEANDTANYCQHS  
 RELPFTFGGGTKVEIKRT (SEQ ID NO:655)

## CD8 transmembrane domain

(DNA)

atctacatctgggcgccttggccgggacttgtgggtccttctcctgtcactgggttatcaccctttactgc (SEQ  
 ID NO:656)

(amino acids)

IYIWAPLAGTCGVLLLSLVITLYC (SEQ ID NO:657)

## 4-1BB domain

(DNA)

aaacggggcagaaagaaactcctgtatatattcaaacaaccatttatgagaccagtacaaactactcaagaggaagat  
 ggctgtagctgccgatttccagaagaagaagaaggaggatgtgaactg (SEQ ID NO:658)

(amino acids)

KRGRKKLLYIFKQPFMRPVQTTQEEDGCSCRFPEEEEGGCEL (SEQ ID NO:659)

## CD3zeta domain

(DNA)

agagtgaagttcagcaggagcgcagacgccccgcgtacaagcagggccagaaccagctctataacgagctcaatcta  
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(amino acids)

RVKFSRSADAPAYKQGQNQLYNELNLGRREEYDVLDRRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMK  
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**CAR-T C2 CD8/CD8/4-1BB/CD3z #44**

N-CD8ls-huMNC2scFv-CD8ecd fragment- CD8 transmembrane- 4-1BB- CD3zeta-C  
(DNA)

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(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKGPGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGT  
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GGSDIVLTQSPASLAVSPGQRATITCRASKSVSTSGYSYMHWYQQKPGQPPKLLIYLASNLESQVPAFSGSGSTG  
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**NFATc1P2-MMP9**

(DNA)

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[illegible]

NFATc1P2-MMP9cat

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#### NFAT response element 2

(DNA)

aagaggaaaatttgtttcatacagaaggcggtt (SEQ ID NO:776)

#### NFAT response element 2 repeats

(DNA)

aagaggaaaatttgtttcatacagaaggcggttaagaggaaaatttgtttcatacagaaggcggttaagaggaaaatttg  
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#### CMV minimal promoter 2

(DNA)

taggcgtgtacggtgggaggcctatataagcagagctcgtttagtgaaccgctcagatcgccctggagacgccatccacg  
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#### NFATRE2mCMV2-MMP9

(DNA)

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#### NFATRE2mCMV2-MMP9cat

(DNA)

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gagtactcgacctgtaccagcgagggcgcgagatgggcgcctctggtgcgctaccacctcgaaactttgacagcgac  
aagaagtgggcttctgcccggaccaaggatacagtttgttctcgtggcgcgcatgagttcgggccacgcgctgggc  
ttagatcattcctcagtgccggagggcgtcatgtacctatgtaccgcttcaactgagggggcccccttgcataaggac  
gacgtgaatggcatccggcacctctatggtcctcgccctgaacctgattacaaggatgacgacgataagtataa  
(SEQ ID NO : 780)

#### NFATc1 Promoter fragment (P1)

(DNA)

aggcaggaggaagaggaaagggcgagggcgctcggggagcagagccgggggcccccggtggccgcagaggccgggc  
cgggcgcgagaggccgggcgagctggccgcgctctgggcgcgcgcctccggaactccctgcgcctggcgcgcgccca  
ccgtggtcccggaacggcattaaacagagggaaacagacccgggattccgtcaccggggcggggggataaggacgg  
ctttgagagcagacaggaagggagcttttctgcatggggtgaaaaattatttattgaaggaggaggaggcgga  
gaggaggaaggggagggggcgaggaggaggaagagccggccgcccccgccccggcctcctcaggagccaag  
ggcagcctcgccaggtcggtcccggtcgaggaccggcgtggggtcgaggggctcagctctccacgtgacgggct  
ggcgcgcccccgccagaccggcctcggtattccctcctcccgcgagctcgcgcgcgcgcctcctggaggtggg  
agaaggaggggcgggcgggggggacggaaactctccccgccaaatcctggccccaggcctggggacactcgcgcg  
gaagatttggaggggaggggagggggagggcggtggggcgcgggcctcgctggagtcctccctgaccccccgacccc  
gcccacggcctggcgctcctcccgggccctcctccctcctcccgcgcccggtgctctggggcgcggtgccacgcct  
ggctcgcgcgctaggggcccccgaggttagagaccctggaaatggcctcgacgcgcgagggagcgaggcgccacc  
acccgcctaactcgggcacgtctctccaggccgagggcctcggtggaaaagccggggttccatttgtgctgagtcgg  
ggcgccgaatggagccaggcctcgggacgcgggacggacgggctctggcgcgccaccttcggggctctgcagcgc  
ccgaccgctccccggcagggaggagggcgttgtggggggcaccacggggcacagtgatccctgggggtctgcgg  
acctcctggggcccccgagcagacacgagtttagcctttgggtttagtttaaatcacataagggtgtcgtgcaatoga  
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tggcagctgagctgcgccgaagagacgcaggagagcgaagagaggaaagtgtgagtgggcgggggcctcccccc  
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ccacgcggccccctgccacagagcgtctaggtcgagcagatatttacagaataaaaaatgacaataactcgacgtcc  
cgggacggccacgcaatctgttagtaatttagcgggatgggaatttcccttctagggcctgccagtgaagcgctttt  
ccaaatttccacagcgggggaagcctgcgattttacataatgaacttcagcatgcgggcttttctgcacacccctccc  
cgggcccccgcccccgcccccgcccccttttccagcagggcggggctcctccggacaccccgctggactcaggcgt  
ccgctctggcccggttcgcccccggtttcccccgccagccccagcgccccctgccccggcccccgattccccgttccc

gccccacgcccccatccccctccccgtgcgccccctccccgtgcgccccccctccccgtgcgccccccctccccgtgcg  
 cccccctccccgtgcgccccccctccccgggcgccccccctccccgggcgccccccctccccgtgcgccccccctcc  
 ccgtgcgccccccctccccgtgcgcccccgctcttgcgccccctgccccaggcgagcggtgcccgcggcgcgggg  
 agggcgggcgctcggcgactcgtccccggggcccccgcggggccccgggcagcagggcgatgacggcagggga  
 gggggcggggagccgccccggcgggggaggcgggggagggtgtttccagctttaaaaaggcaggaggcagagcg  
 cgccctgcgtagagcgagactcagaggctccgaactcgccggcgaggtagcccgcgccagatcccagcagcagggc  
 gcgg (SEQ ID NO:781)

#### NFATc1 Promoter fragment (P2)

(DNA)

aggcaggaggaagaggaaagggcgagggcgctcggggagcagagccgggggccccgggtggccgcagaggccgggc  
 cggggcgagaggccgggcgagctggccgcgctctgggcccgcgcctccggaactccctgcgccctggcgcgcgcca  
 ccgtggtccccggaacggcattaaacagagggaaacagaccgggattccgtcaccggggcggggggataaggacgg  
 ctttgagagcagacaggaaaaggagcttttctgcatggggtgaaaaattatttattgaaggaggaggaggcgga  
 gcggaggaaggggagggcggggaggaggaggaagagccggcccgcccccgcccgccctcaggagcccaag  
 ggcagcctcgccaggtcggtccccgggctcgaggaccggcgctggggctcgaggggctcagctccccacgtgaccggct  
 gggcgcccccgccagaccggcctcggtattccctcctcccgcgagctcctccgcccccgctcctggaggtgggg  
 agaaggaggggcgggggcggggggacggaaactctccccgccaaatcctggccccaggcctggggacactcgcgcg  
 gaagatttgagggggaggggagggggagggcgctggggcgcgccctcgtggagtccccctgacccccgaccccc  
 gccacccggcctggcgctcctcccgcgccccctcctccctcccgcgccccgggtgctctggggcgcgctgccacgct  
 ggctcgcgccgtaggggcccccgaggtagagaccctggaaatggcctcgacggcgaggagcgaggcgccacc  
 acccgctaataccggcacgtctctccaggccgaggcctcggtggaaaagccggggtccatttgctgctgagtcgg  
 ggcgccgaatggagccaggcctcgggacggggacggacgggctctggcgcgccacttcggggctctgcagcgc  
 ccgaccgctcccccgccagggaggaggcgcttgggggggacccacggggcacagtgatccctgggggtctgcgg  
 acctcctgggccccgcagcagacacgagtttagcctttgggttttagtttaaatcacataaggggtgctgcaatcga  
 tttatggtttctacacaccagacactttaacctccaacccccccatccaagccaacaagaaaatgcggtgcccgtgt  
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 gtccagaagtgcgcagctcgccccataaaaacgccccctccggggcggttagggcagggtgagcgcgctccccgggcctcc  
 ccacgcggccccctgccacagagccgtctaggtcgagcagatatttacagaataaaaaatgacaataactcgacgtcc  
 cgggacggccacgcaatctgttagtaatttagcgggatgggaatttcccttctagggcctgccagtgaagcgctttt  
 ccaatttccacagcgggggaagcctgcgattttacataatgacttcagcatgcgggctttctcgacacccctccc  
 cgcccccgcccccgcccccgccccctttccagcaggcgcggtccctcccggaacccccgctggactcaggcgt  
 cccgtctggccgcttcgcccccgtttcccccgccagcccgccccctgccccggccccggattccccggttccc  
 gccccacgcccccatccccctccccgtgcgccccctccccgtgcgccccccctccccgtgcgccccccctccccgtgcg  
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 ccgtgcgccccccctccccgtgcgcccccgctcttgcgccccctgccccaggcgagcggtgcccgcggcgcgggg  
 agggcgggcgctcggcgactcgtccccggggcccccgcggggccccgggcagcagggcgatgacggcagggga  
 gggggcggggagccgccccggcgggggaggcgggggagggtgtttccagctttaaaaaggcaggaggcagagcg  
 cgccctgcgtagagcgagactcagagg (SEQ ID NO:782)

#### NFATc1 Promoter fragment (P3)

(DNA)

caggcctggggacactcgcgcggggaagatttgagggggaggggagggggagggcgctggggggcgcgccctcgctgg  
 agtccccctgaccccccgacccccgcccacggcctggcgctcctcccgcgccccctcctccccctccccggcgccccg  
 tgctctggggcgcgctgccacgctggctcgcgccgttaggggcccccgaggttagagaccctggaaatggcctcga  
 cgccgcaggagcgaggcgccaccaccccgctaataccgggcacgtctctccaggccgaggcctcggtggaaaagcc  
 ggggttccatttgctgctgagtcggggcgccgaatggagccaggcctcgggacggggacggacgggctctggccgc  
 gcaccttcggggctctgcagcgcccgaccgctcccccgccagggaggaggcgcttgggggggacccacggggc  
 acagtatccctgggggtctgcggacctcctgggcccccgagcagacagagtttagcctttgggttagtttaaat  
 cacataaggggtgctgcaatcgatttatggtttctacacaccagacactttaacctccaacccccccatccaaag  
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 gtgagtggccggggggcctcccccgctcagaagtgcgcagctcgccccataaaaacgccccctccggggcggttaggg  
 cagggtgagcgctccccgggcctccccacgcggccccctgccacaggcctctaggtcgagcagatatttacagaat  
 aaaaatgacaataactcgacgtccccgggacggccacgcaatctgttagtaatttagcgggatgggaatttcccttct  
 agggcctgccagtgaagcgcttttccaaatttccacagcggggggaagcctgcgattttacataatgacttcagcatg  
 ccgggctttctcgacacccctccccggcccccgcccccgcccccgcccccttttccagcaggcgcggtccctcc  
 ggacaccccgctggactcaggcgtcccgctctggcccgcttcgccccggtttcccccgccagccccagcggccccctgc

ccggcccccgattccccgttcccgccccctacgcccccatccccctccccgtgccccctccccgtgccccctcc  
 ccgtgccccccccctccccgtgccccccccctccccgtgccccccccctccccggcgccccctccccggcgcccc  
 cctccccgtgccccccccctccccgtgccccccccctccccgtgccccccccctccccgtgccccccccctccccag  
 gcgagcggtgccccggcgcggggagggcgggcgctcggcgactcgccccggggcccccgcggggccccgggcagc  
 agggcggtgatgtcacggcagggagggggcgcgggagcgccccggggcggggagggcggggaggtgtttccagc  
 tttaaaaaggcaggaggcagagcgcgccctgctcagagcgagactcagaggctccgaactcgccggcgaggctgc  
 cgcgccagatcccagcagcagggcgcg (SEQ ID NO:783)

#### **pNFAT-MMP9cat-1 gBLOCK sequence**

(DNA)

aagaggaaaatttgtttcatacagaaggcggttactagttaggcggtgtacgggtgggaggcctatataagcagagctcgt  
 ttagtgaaccgtcagatcgctggagacgccatccacgctgttttgacctccatagaagacaccgggaccgatccagc  
 ctctcgacattcgtttctagagccaccatgagcctctggcagccccctggctcctggtgctcctggtgctgggctgctgc  
 ttgtctttccaaacctttgagggcgacctcaagtggcaccaccacaacatcacctattggatccaaaactactcggaa  
 gacttgccgcgggcggtgattgacgacgcctttgcccgcgccttcgcactgtggagcgcggtgacgcgcgctcaccttc  
 actcgctgtacagccgggacgcagacatcgatccagtttgggtgtcgcgagcacggagacgggtatcccttcgac  
 gggaaggacgggtcctggcacacgcctttcctcctggccccggcattcagggagacgcccatttcgacgatgacgag  
 ttgtggtcctgggcaaggcgctgctggtttccaaactcggtttggaaaacgcagatggcgcgcgctgccacttcccccttc  
 atcttcgagggcgctcctactctgctgcaccaccgacggctcgctcgcagcgttgcctggtgagtagccacggcc  
 aactacgacaccgacgacgggttttggtctctgccccagcgagagactctacaccaggacggcaatgctgatgggaaa  
 cctgcccagtttccattcatcttccaaggccaactcctactcgcctgcaccacggacggctcgctccgacggctaccgc  
 tgggtgcgccaccaccgccaactacgacgggacaagctcttcggcttctgcccgaaccgagctgactcg (SEQ ID  
 NO:784)

#### **NFAT consensus sequence:**

(A/T)GGAAA(A/N)(A/T/C)N (SEQ ID NO:804)

#### **Current NFAT RE (Form System Biosciences. The sequence is from the mouse IL2 promoter)**

(DNA)

aagaggaaaatttgtttcatacagaaggcggtt (SEQ ID NO:805)

#### **Mouse IL2 Promoter (highlighted in green the NFAT RE used, highlighted in yellow is the start codon)**

(DNA)

aactagagacatataaaaataacaccaacatccttagatatacaacccttccctgagaattttattggacatcatactcttt  
 ttaaaaagcataataaacatcaagacacattacacaaaatatgttaaatttaaatttaaaacaacaacgacaaaatagt  
 acctcaagctcaacaagcatttttaggtgtccttagcttactatttctctggttaactgtatgaagccatctatcacc  
 ctgtgtgcaattagctcattgtgtagataagaaggtaaaaccatcttgaaacaggaaaccaatatccttccctgtcta  
 atcaacaaatctaaaagatttattcttttcatctatctcctcttgcggtttgtccaccacaacagggtgcttacaggt  
 tcaggatgggttttgacaaagagaacattttcatgagttacttttgtgtctccaccccaagaggaaaatttgtttca  
 tacagaaggcggttcattgtatgaattaaaactgccacctaagtgtgggctaaccgaccaagagggtattcacctaa  
 atccattcagtcagtgatggtgttttaaaagaaattccagagagtcatacagaagaggaaaaacaaaaggtaatgctt  
 tctgccacacaggtagactctttgaaaatatgtgtaatatgtaaaacatcgtagaccccccatattatttttccagc  
 attaacagtataaattgcctcccatgctgaagagctgctatacccttgctaatcactcctcacagtgaacctcaag  
 tcttcgaggcatgtacagcatgcagctcgcatcctgtgtcac (SEQ ID NO:806)

#### **NFAT RE (Form PRomega. The sequence is from the humane IL2 promoter)**

(DNA)

ggaggaaaaactgtttcatacagaaggcggt (SEQ ID NO:807)

#### **Possible NFAT RE from ET-1 promoter**

(DNA)

tccagggaatcggagtagaacaagagggtatg (SEQ ID NO:808)

#### **Possible NFAT RE from ET-1 promoter**

(DNA)

actgttggaacgtaaacacggttattaaacggt (SEQ ID NO:809)

**Possible NFAT RE from human CD3 $\gamma$**

(DNA)

tccttaacggaaaaacaaaa (SEQ ID NO:810)

**Possible NFAT RE from human CD3 $\gamma$**

(DNA)

aaaggaaaaagtatatgttc (SEQ ID NO:811)

**Possible NFAT RE from human IL3 promoter**

(DNA)

atgccatggaaagggtg (SEQ ID NO:812)

**Possible NFAT RE from human GPC6**

(DNA)

aaggggaaatgttgagtctaga (SEQ ID NO:813)

**Possible NFAT RE from human growth hormone-releasing hormone**

(DNA)

AACTTGGAAGCATAG (SEQ ID NO:814)

**NFATc1 promoter large**

(DNA)

ttatgccgtctagaggagacatactttctactcaaagctacacacatagactacaacgatgggaaaagacgacacac  
caacagcgacttcaggaaagctggagtggtgctgtaaatggttagacaaaataggcttttttaaaaaagggttttattaaag  
aggaatgtttcgtaatgataaaagcactaatctgtgagaaagatacaacaatgataaacatacgtgcagctaataag  
agagctccaaaatctatgaagcaaaaactcacagaatgaggggagaagcagttctacaacagagaatggggacttcg  
atactccactttcaataatggatacaacaaccaggcagataacaaggcaacagaaggcctgaacaacagtataaacc  
aattagacctaccagatatctatagctagcacactccaccaacgacagcagaatacacattcttctcaagcgaca  
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gaaagaaaaagaaaaaaaagtgagcatgtattttgcagagctctggagattagaattaaattagcaaaccagaatt  
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tggttgagctgagaaagaaggatttcccttttagttatgcacctgtgtcagcacctctgactttccttctaaagtc  
tggggtgttctgaggatccgtaagtttgggggttcagggtttctacagcatgctgttacttgtgaaacatctcttta  
accatgtcccagagttgccaggagtttaagaccagcctgagcaacatagcaagacctcatctcaacaacaacaaaa  
attagaaataaattagccagggtgtggtgacatgtgcctgtagtcccagctactcagaaggctgaggcatgaggatca  
cttgggcccaggaagtggggctgcaggagccctgttcagccgctgcactccagcctgcaagacagagcagaaaaa  
aaagaatcaggatcctgggcagagggaggagaggggacccgggtccagcaagcacttggggattgactgaatggcgt  
tggggagagatgactccaaagtccctggagtggtgagaatgactgcgagtggttttaggtggggagggttcctgcct  
ggccactccgggaggggacgtggggctgaagggtatcagggtgccgtgctgagcagtttggccttgatcctaagccc  
tggaacacagcttaggtaggaaagttagctgatccattggtgatctgagtttttagacatggtggtagtccatgag  
gtgggtgttcatgctaagagtttagacagggaaacctatgaagcccttagcaacctccagggaaagggcggtggtta  
aagagatgtttcataagtaacagcatggtatagaaactctgaaccccaaatgtatgggtccctcaggaacaccccaga  
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gctgtgaggttcattcacactgacttaaaaaaaaaaatctcagtttacttaaaagtaaatagctttttctataattc  
tggtttgctaatttaactctagctctccagacctggctaaataaatgcccatttctccagatggtctcaagagttct  
tggaacatcgtgggggccccttccctgttggttggaagggtgctcaggaagaagggggtggttctgagttgagtcaaa  
acctcaaagacccctgatgggaaaagctctcaagtgaaccacgctgtgggcccagaatgcaaaactgcaggaacagaa  
cattcgcaggaacagaacacagctcgtaattaagtgttttcccgagcaggaagtggtcctgacctgcggttcagtag  
gggaggaaagggtgggcgcacctgccctggctgggcgcacctgccaggtagccccacgcggcaccgcgtgtgcga



(DNA)

catcatggggtctgtggaccaaggttaactgactctcgatcccttccagccttttccgctcgctc (SEQ ID NO:818)

**NFATc1 response element consensus**

(DNA)

cattttttccat (SEQ ID NO:819)

**NFATc1 response element consensus**

(DNA)

tttttcca (SEQ ID NO:820)

**NFAT response elements contained within the Foxp3 enhancer region**

(DNA)

acttgaaaatgagataaatgttcacctatgttggtctctagctctcttttatggcttcatt  
ttttccatttactatagagggttaagagtgtgggtactggagccagactgtctgggacaa (SEQ ID NO:821)

N+20

SNIKFRPGSVVVQLTLAFREGTINVHDTVETQFNQYKTEAASRYNLTISDVSVSDVPFPFSAQSGA (SEQ ID NO:822)

N+20/C-27

SNIKFRPGSVVVQLTLAFREGTINVHDTVETQFNQYKTE (SEQ ID NO:823)

N+9/C-9

VQLTLAFREGTINVHDTVETQFNQYKTEAASRYNLTISDVSVSDVP (SEQ ID NO:824)

C-10

GTINVHDTVETQFNQYKTEAASRYNLTISDVSVSDV (SEQ ID NO:825)

NME7-AB

(DNA)

atggaaaaaacgctggccctgattaaaccggatgcaatctccaaagctggcgaaattatcgaaattatcaacaaagcg  
ggtttcaccatcacgaaactgaaaatgatgatgctgagccgtaaagaagccctggattttcatgtcgaccaccagctc  
cgcccggtttttcaatgaactgattcaattcatcaccacgggtccgattatcgcaatggaaattctgcgatgatgacgct  
atctgcgaatggaaacgctgctgggcccggcaaaactcaggtgttgcggtaccgatgccagtgaaatccattcgcgct  
ctgtttggcaccgatggtatccgtaatgcagcacatgggtccggactcattcgcatcggcagctcgtgaaatggaactg  
tttttcccgagctctggcggttgcggtccggcaaacaccgccaaattttaccaattgtacgtgctgtattgtcaaaccg  
cacgcagtgtcagaaggcctgctgggtaaaattctgatggcaatccgtgatgctggctttgaaatctcgccatgcag  
atgttcaacatggaccgcttaacgtcgaagaattctacgaagtttacaaggcggtggttaccgaatatcacgatatg  
gttacggaaatgtactccgggtccgtgctgcgatggaaattcagcaaaacaatgccacaaaacggttctgtgaattc  
tgtgggtccggcagatccggaaatcgacgctcatctgcgtccgggtaccctgcgcgcaatttttggtaaaacgaaaatc  
cagaacgctgtgcactgtaccgatctgccggaagacgggtctgctggaagttcaatactttttcaaaattctggataat  
ctcgagcaccaccaccaccactga (SEQ ID NO:826)

NME7AB

(amino acids)

MEKTLALIKPDAISKAGEIIEIINKAGFTITKLKMMMLSRKEALDFHVDHQSRPFFNELIQFITTGPPIAMEILRDD  
AICEWKRLGPGANSGVARTDASESIRALFGTDGIRNAAHGPDSFASAAREMELFFPSSGGCGPANTAKFTNCTCCIV  
KPHAVSEGLLGKILMAIRDAGFEISAMQMFNMDRVNVEEFYEVYKGVVTEYHDMVTEMYSGPCVAMEIQQNNATKTF  
REFCGPADPEIARHLRPGTLRAIFGKTKIQNAVHCTDLPEDGLLEVQYFFKILDNLEHHHHHH (SEQ ID NO:827)

Human NME7 x1

(DNA)

atgatgatgctttcaaggaaagaagcattggattttcatgtagatcaccagtcagaccctttttcaatgagctgat  
ccagtttattacaactggctctattattgccatggagattttaagagatgatgctatatgtgaatggaaaagactgc

tgggacctgcaaactctggagtggcacgcacagatgcttctgaaagcattagagccctctttggaacagatggcata  
 agaaatgcagcgcattggccctgattcttttgccttctgcggccagagaaatggagttgttttttccctcaagtggagg  
 ttgtgggcccggcaaactgctaaatttactaattgtacctgttgcatgttaaaccatgctgtcagtgaggac  
 tgttgggaaagatcctgatggctatccgagatgcaggttttgaaatctcagctatgcagatgttcaatatggatcgg  
 gttaatgttgaggaattctatgaagtttataaaggagtagtgaccgaatatcatgacatggtgacagaaatgtattc  
 tggcccttgtgtagcaatggagattcaacagaataatgctacaaagacatttcgagaattttgtggacctgctgatc  
 ctgaaattgcccggcatttacgccctggaactctcagagcaatctttggtaaaactaagatccagaatgctgttcac  
 tgtactgatctgccagaggatggcctattagaggttcaatacttcttcaagatcttgataatctcgagcaccacca  
 ccaccaccactga (SEQ ID NO:828)

(amino acids)

MMMLSRKEALDFHVDHQSRPFFNELIQFITTGPPIAMEILRDDAICEWKRLGPNANSGVARTDASESIRALFGTDGIR  
 NAAHGPD SFASAREMELFFPSSGGCGPANTAKFTNCTCCIVKPHAVSEGLLGKILMAIRDAGFEISAMQMFNMDRVN  
 VEEFYEVYKGVVTEYHDMVTEMYSGPCVAMEIQNNATKTFREFCGPADPEIARHLRPGTLRAIFGKTKIQNAVHCTD  
 LPEDGLLEVQYFFKILDNLEHHHHH\* (SEQ ID NO:829)

#### Mouse Antibody 17H6 Heavy chain: DNA sequence

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGTGGCTGAACTGGATTTTCCTTGTAACACTTTTAAATGGTATCCAGTGTGAGGTGAAGCTGGTGGAGTCT  
 GGAGGAGGCTTGGTACAGCCTGGGGGTTCTCTGAGACTCTCCTGTGCAACTTCTGGGTTACCTTCACTGATTACTAC  
 ATGAGCTGGGTCCGCCAGCCTCCAAGAAAGGCACCTTGAGTGGTTGGGTTTTATTAGAAACAAAGCTAATGGTTACACA  
 GCAGAGTACAGTGCCTGTGTGAAGGGTCGGTTCACCATCTCCAGAGATGTTTCCCAAACCTCCTCTATCTTCAAATG  
 AACATCCTGAGAGCTGAGGACAGTGCCACTTATTACTGTGCAAAAGATTACTACGGTAGTAACCTGCCTGGTTTGTCT  
 TACTGGGGCCAAGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:830)

#### Mouse Antibody 17H6 Heavy chain: Amino acid sequence

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLWLNWIFLVTLNLIQCEVKLVESGGGLVQPGGSLRLSCATSGFTFTDYYSWVRQPPRKALEWLGFI RNKANGYT  
 AEYSASVKGRFTISRDSVNLLYLQMNILRAEDSATYYCAKDYYGSNPAWFAYWGQGLVTVSA (SEQ ID  
 NO:831)

Mouse 17H6 heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

GAGGTGAAGCTGGTGGAGTCTGGAGGAGGCTTGGTACAGCCTGGGGGTTCTCTGAGACTCTCCTGTGCAACTTCTGGG  
 TTCACCTTCACT (SEQ ID NO:832)

(amino acids)

EVKLVESGGGLVQPGGSLRLSCATSGFTFT (SEQ ID NO:833)

Mouse 17H6 heavy chain variable *complementarity determining regions 1 (CDR1)*  
 sequence:

(DNA)

GATTACTACATGAGC (SEQ ID NO:834)

(amino acids)

DYYMS (SEQ ID NO:835)

Mouse 17H6 heavy chain variable *framework 2 (FW2)* sequence:

(DNA)

GAGGTGAAGCTGGTGGAGTCTGGAGGAGGCTTGGTACAGCCTGGGGGTTCTCTGAGACTCTCCTGTGCAACTTCTGGG  
 TTCACCTTCACT (SEQ ID NO:836)

(amino acids)

WVRQPPRKALEWLG (SEQ ID NO:837)

Mouse 17H6 heavy chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

TTTATTAGAAACAAAGCTAATGGTTACACAGCAGAGTACAGTGCCTCTGTGAAGGGT (SEQ ID NO:838)

(amino acids)

FIRNKANGYTAEYSASVKG (SEQ ID NO:839)

Mouse 17H6 heavy chain variable *framework 3 (FW3)* sequence:

(DNA)

CGGTTCCACCATCTCCAGAGATGTTTCCCAAACCTCCTCTATCTTCAAATGAACATCCTGAGAGCTGAGGACAGTGCC  
ACTTATTACTGTGCAAAA (SEQ ID NO:840)

(amino acids)

RFTISRDVSQLLYLQMNILRAEDSATYYCAK (SEQ ID NO:841)

Mouse 17H6 heavy chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

GATTACTACGGTAGTAACCCTGCCTGGTTTGCTTAC (SEQ ID NO:842)

(amino acids)

DYYGSNPAWFAY (SEQ ID NO:1)

Mouse 17H6 heavy chain variable *framework 4 (FW4)* sequence:

(DNA)

TGGGGCCAAGGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:843)

(amino acids)

WGQGTLVTVSA (SEQ ID NO:1)

Mouse Antibody **17H6 Light chain: DNA sequence**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGCCTGTGAGGCTGTTGGTGCTGATGTTCTGGATTCCCTGCTTCCAACAGTGATATTTTGATGACCCAGACT  
CCACTCTCCCTGCCTGTCTAGTCTTGGAGATCAAGCCTCCATCTCTTGCAGATCTAGTCAGAGCATTGTACATAGTAGT  
GGAAACACCTTTTTAGAAATGGTACCTGCAGAAACCTGGCCAGTCTCCAAAGCTCCTGATCTACAAAGTTTCCAACCGA  
TTTTCTGGGGTCCCAGACAGGTTTCACTGGCAGTGGATCAGGGATAGATTTTCACTCAAGATCAGCAGAGTGGAGGCT  
GAGGATCTGGGAGTTTATTACTGCTTTCAAGGTTTACATGTTTCTTTACGTTTCGGCTCGGGGACAAAGTTGGAAATA  
AAA (SEQ ID NO:844)

Mouse Antibody **17H6 Light chain: Amino acid sequence**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLPPVRLLVLMFWIPASNSDILMTQTPLSLPVSLGDAQSISCRSSQSIVHSSGNTFLEWYLQKPGQSPKLLIYKVSNR  
FSGVPDRFSGSGSIDFTLKISRVEAEDLGVYYCFQGSHVPFTFGSGTKLEIK (SEQ ID NO:845)

Mouse 17H6 light chain variable *framework 1 (FW1)* sequence:

(DNA)

GATATTTTGATGACCCAGACTCCACTCTCCCTGCCTGTCTAGTCTTGGAGATCAAGCCTCCATCTCTTGC (SEQ ID NO:846)

(amino acids)

DILMTQTPLSLPVSLGDAQSISC (SEQ ID NO:847)

Mouse 17H6 light chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

AGATCTAGTCAGAGCATTGTACATAGTAGTGGAACACCTTTTTAGAA (SEQ ID NO:848)

(amino acids)



RSSQSIVHSSGNTFLE (SEQ ID NO:849)

Mouse 17H6 light chain variable *framework 2 (FW2)* sequence:

(DNA)

TGGTACCTGCAGAAACCTGGCCAGTCTCCAAAGCTCCTGATCTAC (SEQ ID NO:850)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:851)

Mouse 17H6 light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

AAAGTTTCCAACCGATTTTCT (SEQ ID NO:852)

(amino acids)

KVSNRFS (SEQ ID NO:853)

Mouse 17H6 light chain variable *framework 3 (FW3)* sequence:

(DNA)

GGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAGGGATAGATTTTCACTCAAGATCAGCAGAGTGGAGGCTGAGGATCTGGGAGTTTATTACTGC (SEQ ID NO:854)

(amino acids)

GVPDRFSGSGSIDFTLKISRVEAEDLGVYYC (SEQ ID NO:855)

Mouse 17H6 light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

TTTCAAGGTTACATGTTCTTTTACG (SEQ ID NO:856)

(amino acids)

FQGSHPFT (SEQ ID NO:857)

Mouse 17H6 light chain variable *framework 4 (FW4)* sequence:

(DNA)

TTGGGCTCGGGGACAAAGTTGGAAATAAAA (SEQ ID NO:858)

(amino acids)

FGSGTKLEIK (SEQ ID NO:859)

#### **Mouse antibody 39H5 Heavy chain: DNA sequence**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGGCTTGGGTGTGGACCTTGCTATTCCTGATGGCAGCTGCCCAAAGTGCCCAAGCACAGATCCAGTTGGTGCAGTCTGGACCTGAGCTGAAGAAGCCTGGAGAGACAGTCAAGATCTCCTGCAAGGCTTCTGGGTATACCTTCACAACTATGGAATGAAGTGGGTGAAGCAGGCTCCAGGAAAGGGTTTAAAGTGGATGGGCTGGATAAACACCTACACTGGAGAGCCAACA TATGTTGGTGACTTCAAGGGACGGTTTGCCTTCTCTTTGGAGACCTCTGCCAGCACTGCCTATTTGCAGATCAACAACCTCAAAAATGAGGACACGGCTACATATTTTGTGTTAGAGGTATCCACGGCTACGTGGACTACTGGGGCCAAGGCACC ACTCTCACAGTCTCCTCA (SEQ ID NO:860)

#### **Mouse antibody 39H5 Heavy chain: Amino acid sequence**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MAWVWTLFLMAAAQSAQAQIQLVQSGPELKKPGETVKISKASGYTFTNYGMNWVKQAPGKGLKWMGWINTYTGEPT YVGDFKGRFAFSLETSASTAYLQINNLLKNEDTATYFCVRGIHGYYVDYWGQGTTLTVSS (SEQ ID NO:861)

**Mouse antibody 39H5 heavy chain variable *framework 1 (FW1)* sequence:**

(DNA)

CAGATCCAGTTGGTGCAGTCTGGACCTGAGCTGAAGAAGCCTGGAGAGACAGTCAAGATCTCCTGCAAGGCTTCTGGG  
TATACCTTCACA (SEQ ID NO:862)  
(amino acids)

QIQLVQSGPELKKPGETVKISCKASGYTFT (SEQ ID NO:863)

**Mouse antibody 39H5 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AACTATGGAATGAAC (SEQ ID NO:864)

(amino acids)

NYGMN (SEQ ID NO:865)

**Mouse antibody 39H5 heavy chain variable framework 2 (FW2) sequence:**

(DNA)

TGGGTGAAGCAGGCTCCAGGAAAGGGTTTAAAGTGGATGGGC (SEQ ID NO:866)

(amino acids)

WVKQAPGKGLKWMG (SEQ ID NO:867)

**Mouse antibody 39H5 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

TGGATAAACACCTACACTGGAGAGCCAACATATGTTGGTGACTIONCAAGGGA (SEQ ID NO:868)

(amino acids)

WINTYTGEPTYVGDFKG (SEQ ID NO:869)

**Mouse antibody 39H5 heavy chain variable framework 3 (FW3) sequence:**

(DNA)

CGGTTTGCCTTCTCTTTGGAGACCTCTGCCAGCACTGCCTATTTGCAGATCAACAACCTCAAAAATGAGGACACGGCT  
ACATATTTTTGTGTTAGA (SEQ ID NO:870)

(amino acids)

RFAFSLETSASTAYLQINNLLKNETATYFCVR (SEQ ID NO:871)

**Mouse antibody 39H5 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

GGTATCCACGGCTACGTGGACTAC (SEQ ID NO:872)

(amino acids)

GIHGYVDY (SEQ ID NO:873)

**Mouse antibody 39H5 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGCCAAGGCACCACTCTCACAGTCTCCTCA (SEQ ID NO:874)

(amino acids)

WGQGTTLTVSS (SEQ ID NO:875)

**Mouse antibody 39H5 Light chain: DNA sequence**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGCCTGTTAGGCTGTTGGTGCTGATGTTCTGGATTCTCCTTCCAGCAGTGATGTTTTGATGACCCAAACT  
CCACTCTCCCTGCCTGTCTAGTCTTGGAGATCAAGCCTCCATCTCTTGCAGATCTAGTCAGAGCATTGTACATAGAAAT  
GGAAACACCTATTTAGAATGGTACCTGCAGAAACCAGGCCAGTCTCCAAAGCTCCTGATCTACAAAGTTTCCAACCGA  
TTTTCTGGGGTCCCAGACAGGTTCACTGGCAGTGGATCAGGGACAGATTTCACTCAAGATCAGCAGAGTGGAGGCT  
GAGGATCTGGGAGTTTATTACTGCTTTCAAGGTTACATCTTCCGTGGACGTTCCGGTGGAGGCACCAAGCTGGAAATC  
AAA (SEQ ID NO:876)

**Mouse antibody 39H5 Light chain: Amino acid sequence**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLPVRLLVLMFWIPASSSDVLTQTPLSLPVS LGDQASISCRSSQSIVHRNGNTYLEWYLQKPGQSPKLLIYKVS NR  
FSGVPDRFSGSGSGTDFTLKISRVEAEDLGVYYCFQGSHLPWTFGGGTKLEIK (SEQ ID NO:877)

**Mouse antibody 39H5** light chain variable *framework 1 (FW1)* sequence:

(DNA)

GATGTTTTGATGACCCAACTCCACTCTCCCTGCCTGTCAGTCTTGGAGATCAAGCCTCCATCTCTTGC (SEQ ID NO:878)

(amino acids)

DVLTMTQTPLSLPVS LGDQASIS (SEQ ID NO:879)

**Mouse antibody 39H5** light chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

AGATCTAGTCAGAGCATTGTACATAGAAATGGAAACACCTATTTAGAA (SEQ ID NO:880)

(amino acids)

RSSQSIVHRNGNTYLE (SEQ ID NO:881)

**Mouse antibody 39H5** light chain variable *framework 2 (FW2)* sequence:

(DNA)

TGGTACCTGCAGAAACCAGGCCAGTCTCCAAAGCTCCTGATCTAC (SEQ ID NO:882)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:883)

**Mouse antibody 39H5** light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

AAAGTTTCCAACCGATTTTCT (SEQ ID NO:884)

(amino acids)

KVSNRFS (SEQ ID NO:885)

**Mouse antibody 39H5** light chain variable *framework 3 (FW3)* sequence:

(DNA)

GGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAGGGACAGATTTTCACTCAAGATCAGCAGAGTGGAGGCTGAGGAT  
CTGGGAGTTTATTACTGC (SEQ ID NO:886)

(amino acids)

GVPDRFSGSGSGTDFTLKISRVEAEDLGVYYC (SEQ ID NO:887)

**Mouse antibody 39H5** light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

TTTCAAGGTTACATCTTCCGTGGACG (SEQ ID NO:888)

(amino acids)

FQGSHPWT (SEQ ID NO:889)

**Mouse antibody 39H5** light chain variable *framework 4 (FW4)* sequence:

(DNA)

TTCGGTGGAGGCACCAAGCTGGAAATCAAA (SEQ ID NO:890)

(amino acids)

FGGGTKLEIK (SEQ ID NO:891)

**Antibody 3C5 Heavy chain: DNA sequence**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGGCTTGGGTGTGGACCTTGCTGTTCTTGATGGCAGCTGCCCAAAGTGCCCAAGCACAGATCCAGTTGGTGCAGTCT  
GGACCTGAGCTGAAGAAGCCTGGAGAGACAGTCAAGATCTCCTGCAAGGCTTCTGGGTATACCTTCACAACTATGGA  
ATGAAGTGGGTGAAGCAGGCTCCAGGAAAGGGTTTAAAGTGGATGGGCTGGATAAACACCTACACTGGAAAGCCAACA

TATGCTGATGACTTCAAGGGACGGTTTGCCTTCTCTTTGGAGACCTCTGCCAGCACTGCCTATTTGCAGATCAACAAC  
CTCAAAAATGAGGACACGGCTACATATTTCTGTGCAAGAGGGGGACTAGATGGTTACTACGGCTACTGGGGCCAAGGC  
ACCACTCTCACAGTCTCCTCA (SEQ ID NO:892)

**Antibody 3C5 Heavy chain: Amino acid sequence**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MAWVWTLFLMAAAQSAQAQIQLVQSGPELKKPGETVKISKASGYTFTNYGMNWVKQAPGKGLKWMGWINTYTCKPT  
YADDFKGRFAFSLETSASTAYLQINNLLKNETATYFCARGGLDGYGYWGQGTTLTVSS (SEQ ID NO:893)

**Mouse antibody 3C5 heavy chain variable framework 1 (FW1) sequence:**

(DNA)

CAGATCCAGTTGGTGCAGTCTGGACCTGAGCTGAAGAAGCCTGGAGAGACAGTCAAGATCTCCTGCAAGGCTTCTGGG  
TATACCTTCACA (SEQ ID NO:894)

(amino acids)

QIQLVQSGPELKKPGETVKISKASGYTFT (SEQ ID NO:895)

**Mouse antibody 3C5 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AACTATGGAATGAAC (SEQ ID NO:896)

(amino acids)

NYGMN (SEQ ID NO:897)

**Mouse antibody 3C5 heavy chain variable framework 2 (FW2) sequence:**

(DNA)

TGGGTGAAGCAGGCTCCAGGAAAGGGTTTAAAGTGGATGGGC (SEQ ID NO:898)

(amino acids)

WVKQAPGKGLKWMG (SEQ ID NO:899)

**Mouse antibody 3C5 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

TGGATAAACACCTACACTGGAAAGCCAACATATGCTGATGACTTCAAGGGA (SEQ ID NO:900)

(amino acids)

WINTYTCKPTYADDFKG (SEQ ID NO:901)

**Mouse antibody 3C5 heavy chain variable framework 3 (FW3) sequence:**

(DNA)

CGGTTTGCCTTCTCTTTGGAGACCTCTGCCAGCACTGCCTATTTGCAGATCAACAACCTCAAAAATGAGGACACGGCT  
ACATATTTCTGTGCAAGA (SEQ ID NO:902)

(amino acids)

RFAFSLETSASTAYLQINNLLKNETATYFCAR (SEQ ID NO:903)

**Mouse antibody 3C5 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

GGGGGACTAGATGGTTACTACGGCTAC (SEQ ID NO:904)

(amino acids)

GGLDGYGY (SEQ ID NO:905)

**Mouse antibody 3C5 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGCCAAGGCACCACTCTCACAGTCTCCTCA (SEQ ID NO:906)

(amino acids)

WGQGTTLTVSS (SEQ ID NO:907)

**Antibody 3C5 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAGTCCTGCCCAGTTCCTGTTTCTGCTAGTGCTCTCGATTTCAGGAAACCAACGGTGATGTTGTGATGGCTCAGACC  
CCACTCACTTTGTCGGTTACCATTGGACAACCAGCCTCCATCTCTTGCAAATCAAGTCAGAGCCTCTTACATAGTAAA  
GGAAAGACATATTTGAATTGGTTATTACAGAGGCCAGGCCAGTCTCCAAAGCTCCTAATCTATCTGGTGTCTAAACTG  
GAATCTGGAGTCCCTGACAGGTTCACTGGCAGTGGATCAGGGACAGATTTACACTGAAAATCAGCAGAGTGGAGGCT  
GAAGATTTGGGAGTTTATTACTGCTTGCAAACCTACACATTTTCCGTGGACGTTCCGGTGGAGGCACCAAGCTGGAAATC  
AAA (SEQ ID NO:908)

**Antibody 3C5 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MSPAQFLFLVLVLSIQETNGDVVMAQTPLTSLVTIGQPASISCKSSQSLHSHKSKTYLNLWLLQRPQGSPKLLIYLVSKL  
ESGVPDRFSGSGSGTDFTLKISRVEAEDLGVYYCLQTTHFPWTFGGGKLEIK (SEQ ID NO:909)

**Mouse antibody 3C5 light chain variable framework 1 (FW1) sequence:**

(DNA)

GATGTTGTGATGGCTCAGACCCCACTCACTTTGTCGGTTACCATTGGACAACCAGCCTCCATCTCTTGC (SEQ ID NO:910)

(amino acids)

DVMAQTPLTSLVTIGQPASISC (SEQ ID NO:911)

**Mouse antibody 39H5 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AAATCAAGTCAGAGCCTCTTACATAGTAAAGGAAAGACATATTTGAAT (SEQ ID NO:912)

(amino acids)

KSSQSLHSHKSKTYLN (SEQ ID NO:913)

**Mouse antibody 3C5 light chain variable framework 2 (FW2) sequence:**

(DNA)

TGGTTATTACAGAGGCCAGGCCAGTCTCCAAAGCTCCTAATCTAT (SEQ ID NO:914)

(amino acids)

WLLQRPQGSPKLLIY (SEQ ID NO:915)

**Mouse antibody 3C5 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

CTGGTGTCTAAACTGGAATCT (SEQ ID NO:916)

(amino acids)

LVSKLES (SEQ ID NO:917)

**Mouse antibody 3C5 light chain variable framework 3 (FW3) sequence:**

(DNA)

GGAGTCCCTGACAGGTTCACTGGCAGTGGATCAGGGACAGATTTACACTGAAAATCAGCAGAGTGGAGGCTGAAGAT  
TTGGGAGTTTATTACTGC (SEQ ID NO:918)

(amino acids)

GVPDRFSGSGSGTDFTLKISRVEAEDLGVYYC (SEQ ID NO:919)

**Mouse antibody 3C5 light chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

TTGCAAACCTACACATTTTCCGTGGACG (SEQ ID NO:920)

(amino acids)

LQTTHFPWT (SEQ ID NO:921)

**Mouse antibody 3C5 light chain variable framework 4 (FW4) sequence:**

(DNA)

TTCCGTGGAGGCACCAAGCTGGAAATCAAA (SEQ ID NO:922)

(amino acids)

FGGGTKLEIK (SEQ ID NO:923)

**Mouse antibody 8A9 Heavy chain: DNA sequence (420 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGTGGCTGAACTGGATTTCCTTGTAACACTTTTAAATGGTATCCAGTGTGAGGTGGAGCTGGTGGAGTCT  
GGAGGAGGCTTGGTACAGCCTGGGGGTTCTCTGAGACTCTCCTGTGCAACTTCTGGGTTACCTTCACTGATCACTAC  
ATGAGCTGGGTCCGCCAGCCTCCAGGAAAGGCACCTTGAGTGGTTGGGATTTATTAGAAACAAAGCTAATGGTTACACA  
ACAGAGTACAGTGCATCTGTGAAGGGTCGGTTCACCATCTCCAGAGATAATTCCCAAAGCATCCTCTATCTTCAAATG  
AAAACCTGAGAACTGAGGACAGTGCCACTTATTACTGTGCAAGACCTTCTGACTGGGACTCCTGGTTTGCTTACTGG  
GGCCAAGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:924)

**Mouse antibody 8A9 Heavy chain: Amino acid sequence (140 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLWLNWIFLVTLNIGIQCELVESGGGLVQPGGSLRLSCATSGFTFTDHYMSWVRQPPGKALEWLGFI RNKANGYT  
TEYSASVKGRFTISRDNQSILYLQMKTLRTEDSATYYCARPSDWD SWFAYWGQGLVTVSA (SEQ ID NO:925)

**Mouse antibody 8A9 heavy chain variable framework 1 (FW1) sequence:**

(DNA)

GAGGTGGAGCTGGTGGAGTCTGGAGGAGGCTTGGTACAGCCTGGGGGTTCTCTGAGACTCTCCTGTGCAACTTCTGGG  
TTCACCTTCACT (SEQ ID NO:926)

(amino acids)

EVELVESGGGLVQPGGSLRLSCATSGFTFT (SEQ ID NO:927)

**Mouse antibody 8A9 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

GATCACTACATGAGC (SEQ ID NO:928)

(amino acids)

DHYMS (SEQ ID NO:929)

**Mouse antibody 8A9 heavy chain variable framework 2 (FW2) sequence:**

(DNA)

TGGGTCCGCCAGCCTCCAGGAAAGGCACCTTGAGTGGTTGGGA (SEQ ID NO:930)

(amino acids)

WVRQPPGKALEWLG (SEQ ID NO:931)

**Mouse antibody 8A9 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

TTTATTAGAAACAAAGCTAATGGTTACACAACAGAGTACAGTGCATCTGTGAAGGGT (SEQ ID NO:932)

(amino acids)

FIRNKANGYTTEYSASVKG (SEQ ID NO:933)

**Mouse antibody 8A9 heavy chain variable framework 3 (FW3) sequence:**

(DNA)

CGGTTACCATCTCCAGAGATAATTCCCAAAGCATCCTCTATCTTCAAATGAAAACCTGAGAACTGAGGACAGTGCC  
ACTTATTACTGTGCAAGA (SEQ ID NO:934)

(amino acids)

RFTISRDNQSILYLQMKTLRTEDSATYYCAR (SEQ ID NO:935)

**Mouse antibody 8A9 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

CCTTCTGACTGGGACTCCTGGTTTGCTTAC (SEQ ID NO:936)

(amino acids)

PSDWD SWFAY (SEQ ID NO:937)

**Mouse antibody 8A9 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGCCAAGGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:938)

(amino acids)

WGQGTLLVTVSA (SEQ ID NO:939)

**Mouse antibody 8A9 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGCCTGTTAGGCTGTTGGTGCTGATGTTCTGGATTCTGCTTCCAGCAGTGATGTTTTGATGACCCAAACT  
 CCACTCTCCCTGCCTGTCAGTCTTGGTGATCAAGCCTCCATCTCTTGCAGATCTAGTCAGAGCATTGTACATAGTAAT  
 GGCAACACCTATTTAGATTGGTACTTGCAGAAACCAGGCCAGTCTCCAAAGCTCCTGATCTACAGAGTTTCCAACCGA  
 TTTTCTGGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAGGGACAGATTTCACTCAAGATCAGCAGAGTGGAGGCT  
 GAGGATCTGGGACTTTATTACTGTTTTCAAGGTTTCACATGTTCCGTGGGCGTTCGGTGGAGGCACCAAGCTGGAAATC  
 AAA (SEQ ID NO:940)

**Mouse antibody 8A9 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLPVRLLVLMFWIPASSSDVLMQTPLSLPVS LGDQASISCRSSQSIVHSNGNTYLDWYLQKPGQSPKLLIYRVSNR  
 FSGVPDRFSGSGSGTDFTLKISRVEAEDLGLYYCFQGSHPVWAFGGGTKLEIK (SEQ ID NO:941)

**Mouse antibody 8A9 light chain variable framework 1 (FW1) sequence:**

(DNA)

GATGTTTTGATGACCCAAACTCCACTCTCCCTGCCTGTCAGTCTTGGTGATCAAGCCTCCATCTCTTGC (SEQ ID NO:942)

(amino acids)

DVLMTQTPLSLPVS LGDQASISC (SEQ ID NO:943)

**Mouse antibody 8A9 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AGATCTAGTCAGAGCATTGTACATAGTAATGGCAACACCTATTTAGAT (SEQ ID NO:944)

(amino acids)

RSSQSIVHSNGNTYLD (SEQ ID NO:945)

**Mouse antibody 8A9 light chain variable framework 2 (FW2) sequence:**

(DNA)

TGGTACTTGCAAGAACAGGCCAGTCTCCAAAGCTCCTGATCTAC (SEQ ID NO:946)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:947)

**Mouse antibody 8A9 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

AGAGTTTCCAACCGATTTTCT (SEQ ID NO:948)

(amino acids)

RVS NRFS (SEQ ID NO:949)

**Mouse antibody 8A9 light chain variable framework 3 (FW3) sequence:**

(DNA)

GGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAGGGACAGATTTCACTCAAGATCAGCAGAGTGGAGGCTGAGGAT  
 CTGGGACTTTATTACTGT (SEQ ID NO:950)

(amino acids)

GVPDRFSGSGSGTDFTLKISRVEAEDLGLYYC (SEQ ID NO:951)

**Mouse antibody 8A9 light chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

TTTCAAGGTTTCACATGTTCCGTGGGCG (SEQ ID NO:952)

(amino acids)  
FQGSHPWA (SEQ ID NO:953)

**Mouse antibody 8A9** light chain variable *framework 4 (FW4)* sequence:

(DNA)  
TTCGGTGGAGGCACCAAGCTGGAAATCAAA (SEQ ID NO:954)  
(amino acids)  
FGGGTKLEIK (SEQ ID NO:955)

**Mouse antibody 18G12 Heavy chain: DNA sequence (399 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4  
ATGGGATGGAGCTATATCATCCTCTTTTTGGTCGCAACAGCTACAGGTGTCCACTCCCAGGTCCAACCTGCAGCAGTCT  
GGGGCTGAACTGGTGAAGCCTGGGGCTTCAGTGAAGTTGTCTCTGCAAGGCTTCTGGCTACACCTTCACCGGCTACTTT  
TTGTACTGGGTGAAGCAGAGGCCTGGACAAGGCCTTGAGTGGATTGGGGGGATTAATCCTGACAATGGTGGTATTGAC  
TTCAATGAGAAGTTCAGGAACAAGGCCACACTGACTGTAGACAAATCCTCCAGCACAGCCTACATGCAACTCAGCAGC  
CTGACATCTGAGGACTCTGCGGTCTATTATTGTACATTACTAATAGGGAACCTATTGGGGCCAAGGCACCACTCTCACA  
GTCTCCTCA (SEQ ID NO:956)

**Mouse antibody 18G12 Heavy chain: Amino acid sequence (133 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4  
MGWSYIIILFLVATATGVHSQVQLQQSGAELVKPGASVKLSCKASGYTFTGYFLYWVKQRPQGQLEWIGGINPDNGGID  
FNEKFRNKATLTVDKSSSTAYMQLSSLTSEDSAVYYCTLLIGNYWQGTTTLTVSS (SEQ ID NO:957)

**Mouse antibody 18G12** heavy chain variable *framework 1 (FW1)* sequence:

(DNA)  
CAGGTCCAACCTGCAGCAGTCTGGGGCTGAACTGGTGAAGCCTGGGGCTTCAGTGAAGTTGTCTCTGCAAGGCTTCTGGC  
TACACCTTCACC (SEQ ID NO:958)  
(amino acids)  
QVQLQQSGAELVKPGASVKLSCKASGYTFT (SEQ ID NO:959)

**Mouse antibody 39H5** heavy chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)  
GGCTACTTTTTGTAC (SEQ ID NO:960)  
(amino acids)  
GYFLY (SEQ ID NO:961)

**Mouse antibody 18G12** heavy chain variable *framework 2 (FW2)* sequence:

(DNA)  
TGGGTGAAGCAGAGGCCTGGACAAGGCCTTGAGTGGATTGGG (SEQ ID NO:962)  
(amino acids)  
WVKQRPQGQLEWIG (SEQ ID NO:963)

**Mouse antibody 18G12** heavy chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)  
GGGATTAATCCTGACAATGGTGGTATTGACTTCAATGAGAAGTTCAGGAAC (SEQ ID NO:964)  
(amino acids)  
GINPDNGGIDFNEKFRN (SEQ ID NO:965)

**Mouse antibody 18G12** heavy chain variable *framework 3 (FW3)* sequence:

(DNA)  
AAGGCCACACTGACTGTAGACAAATCCTCCAGCACAGCCTACATGCAACTCAGCAGCCTGACATCTGAGGACTCTGCG  
GTCTATTATTGTACATTA (SEQ ID NO:966)  
(amino acids)  
KATLTVDKSSSTAYMQLSSLTSEDSAVYYCTL (SEQ ID NO:967)



**Mouse antibody 18G12 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

CTAATAGGGAAGTAT (SEQ ID NO:968)

(amino acids)

LIGNY (SEQ ID NO:969)

**Mouse antibody 18G12 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGCCAAGGCACCACTCTCACAGTCTCCTCA (SEQ ID NO:970)

(amino acids)

WGQGTTLTVSS (SEQ ID NO:971)

**Mouse antibody 18G12 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAGTCCTGCCAGTTCCTGTTTCTGTTAGTGCTCTGGATTTCGGGAAACCAATGGTGATGTTGTGATGACCCAGACT  
 CCACTCACTTTGTGCGTAACCAATTGGACAGCCAGCCTCCATCTCTTGCAAGTCAAGTCAGAGCCTCTTACATAGTGAT  
 GGAAAGACATATTTGATTGTTGTTTACAGAGGCCAGGCCAGTCTCCAAAGCGCCTAATCTATCTGGTGTCTAAACTG  
 GACTCTGGAGTCCCTGACAGGTTCACTGGCAGTGGATCAGGGACAGATTTACACTGAAAATCAGCAGAGTGGAGGCT  
 GAGGATTTGGGAGTTTATTTTTGCTGTCAAGGTACACATTTTCCGTGGACGTTTCGGTGGAGGCACCATGCTGGAAATC  
 AAA (SEQ ID NO:972)

**Mouse antibody 18G12 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MSPAQFLFLLVLWIRETNGDVMTQTPLTSLVTIGQPASISCKSSQSLLHSDGKTYLIWLLQRPQGSPKRLIYLVSKL  
 DSGVPDRFTGSGSGTDFTLKISRVEAEDLGVIYFCCQGTGTFPWTFGGGTMLEIK (SEQ ID NO:973)

**Mouse antibody 18G12 light chain variable framework 1 (FW1) sequence:**

(DNA)

GATGTTGTGATGACCCAGACTCCACTCACTTTGTGCGTAACCAATTGGACAGCCAGCCTCCATCTCTTGC (SEQ ID  
 NO:974)

(amino acids)

DVVMTQTPLTSLVTIGQPASISC (SEQ ID NO:975)

**Mouse antibody 18G12 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AAGTCAAGTCAGAGCCTCTTACATAGTGATGGAAAGACATATTTGATT (SEQ ID NO:976)

(amino acids)

KSSQSLLHSDGKTYLI (SEQ ID NO:977)

**Mouse antibody 18G12 light chain variable framework 2 (FW2) sequence:**

(DNA)

TGGTTGTTACAGAGGCCAGGCCAGTCTCCAAAGCGCCTAATCTAT (SEQ ID NO:978)

(amino acids)

WLLQRPQGSPKRLIY (SEQ ID NO:979)

**Mouse antibody 18G12 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

CTGGTGTCTAAACTGGACTCT (SEQ ID NO:980)

(amino acids)

LVSKLDS (SEQ ID NO:981)

**Mouse antibody 18G12 light chain variable framework 3 (FW3) sequence:**

(DNA)

GGAGTCCCTGACAGGTTCACTGGCAGTGGATCAGGGACAGATTTCACTGAAAATCAGCAGAGTGGAGGCTGAGGAT  
TTGGGAGTTTATTTTGC (SEQ ID NO:982)

(amino acids)

GVPDRFTGSGSGTDFTLKISRVEAEDLGVYFC (SEQ ID NO:983)

**Mouse antibody 18G12 light chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

TGTCAAGGTACACATTTTCCGTGGACG (SEQ ID NO:984)

(amino acids)

CQGTHFPWT (SEQ ID NO:985)

**Mouse antibody 18G12 light chain variable framework 4 (FW4) sequence:**

(DNA)

TTCGGTGGAGGCACCATGCTGGAAATCAAA (SEQ ID NO:986)

(amino acids)

FGGGTMLEIK (SEQ ID NO:987)

**Mouse antibody 20A10 Heavy chain: DNA sequence (417 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAACCTCGGGTTCAGCTTGATTTTCCTTGTCTTGTGTTTAAAGGTGTCCAGTGTGAAGTGATGCTGGTGGAGTCT  
GGGGGAGGCTTAGTGAAGCCTGGAGGGTCCCTGAACTCTCCTGTGCAGCCTCTGGATTCACTTTCAGTACCTATGCCA  
TGTCTTGGATTGCGCCAGACTCCAGAGAAGAGGCTGGAGTGGGTCCGATCCATTGGTCGTGCTGGTTCCACCTACTATT  
CAGACAGTGTGAAGGGCCGATTACCATCTCCAGAGATAATGTCCGGAACATCCTGTACCTGCAAATGAGCAGTCTGA  
GGTCTGAGGACACGGCCATGTATTACTGTGCTAGAGGCCGATCTACAATGATTACGACGAGTTTGCTTACTGGGGCC  
AAGGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:988)

**Mouse antibody 20A10 Heavy chain: Amino acid sequence (139 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MNFGFSLIFLVLVLKGVQCEVMLVESGGGLVKPGGSLKLSCAASGFTFSTYAMSWIRQTPEKRLEWVASIGRAGSTYY  
SDSVKGRFTISRDNVRNILYLQMSSLRSEDTAMYYCARGPIYNDYDEFAYWGQGLVTVSA (SEQ ID NO:989)

**Mouse antibody 20A10 heavy chain variable framework 1 (FW1) sequence:**

(DNA)

GAAGTGATGCTGGTGGAGTCTGGGGGAGGCTTAGTGAAGCCTGGAGGGTCCCTGAACTCTCCTGTGCAGCCTCTGGA  
TTCACCTTTCAGT (SEQ ID NO:990)

(amino acids)

EVMLVESGGGLVKPGGSLKLSCAASGFTFS (SEQ ID NO:991)

**Mouse antibody 20A10 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

ACCTATGCCATGTCT (SEQ ID NO:992)

(amino acids)

TYAMS (SEQ ID NO:993)

**Mouse antibody 20A10 heavy chain variable framework 2 (FW2) sequence:**

(DNA)

TGGATTGCCAGACTCCAGAGAAGAGGCTGGAGTGGGTCGCA (SEQ ID NO:994)

(amino acids)

WIRQTPEKRLEWVA (SEQ ID NO:995)

**Mouse antibody 20A10 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

TCCATTGGTCGTGCTGGTTCCACCTACTATTTCAGACAGTGTGAAGGGC (SEQ ID NO:996)

(amino acids)

SIGRAGSTYYSDSVKG (SEQ ID NO:997)

**Mouse antibody 20A10 heavy chain variable framework 3 (FW3) sequence:**

(DNA)

CGATTACCATCTCCAGAGATAATGTCCGGAACATCCTGTACCTGCAAATGAGCAGTCTGAGGTCTGAGGACACGGCC  
ATGTATTACTGTGCTAGA (SEQ ID NO:998)

(amino acids)

RFTISRDNVRNILYLQMSSLRSED TAMYYCAR (SEQ ID NO:999)

**Mouse antibody 20A10 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

GGCCCCGATCTACAATGATTACGACGAGTTTGCTTAC (SEQ ID NO:1000)

(amino acids)

GPIYNDYDEFAY (SEQ ID NO:1001)

**Mouse antibody 20A10 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGCCAAGGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:1002)

(amino acids)

WGQGTLLTVSA (SEQ ID NO:1003)

**Mouse antibody 20A10 Light chain: DNA sequence (396 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGGAATCACAGACTCAGGTCTTCTCTCCCTGCTGCTCTGGGTATCTGGTACCTGTGGGAACATTATGATGACACAG  
TCGCCATCATCTCTGGCTGTGTCTGCAGGAGAAAAGGTCATATGAGCTGTAAGTCCAGTCAAAGTGTTTTATACAGT  
TCAAATCAGAAGAACTATTTGGCCTGGTACCAGCAGAAACCAGGGCAGTCTCCTAAACTGCTGATCTACTGGGCATCC  
ACTAGGGAATCTGGTGTCCCTGATCGCTTCACAGGCAGTGGATCTGGGACAGATTTTACTCTTACCATCAGCAGTGTA  
CAAGCTGAAGACCTGGCAGTTTATTACTGTTCATCAATACCTCTCCTCGCTCACGTTCCGGTGCTGGGACCAAGCTGGAG  
CTGAAA (SEQ ID NO:1004)

**Mouse antibody 20A10 Light chain: Amino acid sequence (132 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MESQTQVFLSLLLWVSGTCGNIMMTQSPSSLAVSAGEKVTMCKSSQSVLYSSNQKNYLAWYQQKPGQSPKLLIYWAS  
TRESGVPDRFTGSGSGTDFLTITISSVQAEDLAVYYCHQYLSSLTFGAGTKLELK (SEQ ID NO:1005)

**Mouse antibody 20A10 light chain variable framework 1 (FW1) sequence:**

(DNA)

AACATTATGATGACACAGTCGCCATCATCTCTGGCTGTGTCTGCAGGAGAAAAGGTCATATGAGCTGT (SEQ ID  
NO:1006)

(amino acids)

NIMMTQSPSSLAVSAGEKVTMSC (SEQ ID NO:1007)

**Mouse antibody 20A10** light chain variable *complementarity determining regions* 1 (*CDR1*) sequence:

(DNA)

AAGTCCAGTCAAAGTGTTTTATACAGTTCAAATCAGAAGAACTATTTGGCC (SEQ ID NO:1008)

(amino acids)

KSSQSVLYSSNQKNYLA (SEQ ID NO:1009)

**Mouse antibody 20A10** light chain variable *framework 2 (FW2)* sequence:

(DNA)

TGGTACCAGCAGAAACCAGGGCAGTCTCCTAAACTGCTGATCTAC (SEQ ID NO:1010)

(amino acids)

WYQQKPGQSPKLLIY (SEQ ID NO:1011)

**Mouse antibody 20A10** light chain variable *complementarity determining regions* 2 (*CDR2*) sequence:

(DNA)

TGGGCATCCACTAGGGAATCT (SEQ ID NO:1012)

(amino acids)

WASTRES (SEQ ID NO:1013)

**Mouse antibody 20A10** light chain variable *framework 3 (FW3)* sequence:

(DNA)

GGTGTCCCTGATCGCTTCACAGGCAGTGGATCTGGGACAGATTTTACTCTTACCATCAGCAGTGTACAAGCTGAAGAC  
CTGGCAGTTTATTACTGT (SEQ ID NO:1014)

(amino acids)

GVPDRFTGSGSGTDFLTITSSVQAEDLAVYYC (SEQ ID NO:1015)

**Mouse antibody 20A10** light chain variable *complementarity determining regions* 3 (*CDR3*) sequence:

(DNA)

CATCAATACCTCTCCTCGCTCACG (SEQ ID NO:1016)

(amino acids)

HQYLSSLT (SEQ ID NO:1017)

**Mouse antibody 20A10** light chain variable *framework 4 (FW4)* sequence:

(DNA)

TTCGGTGCTGGGACCAAGCTGGAGCTGAAA (SEQ ID NO:1018)

(amino acids)

FGAGTKLELK (SEQ ID NO:1019)

**Mouse antibody 25E6 Heavy chain: DNA sequence (414 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAACTTCGGGCTCAGCTTGATTTTCCTTGCCCTCATTTTAAAGGTGTCCAGTGTGAGGTGCAGCTGGTGGAGTCT  
GGGGGAGACTTAGTGAAGCCTGGAGGGTCCCTGAACTCTCCTGTGCAGCCTCTGGTTTCACTTTCACTAGTTATGGA  
ATGTCTTGGGTTCCGACAGCTCCAGACAAGAGGCTGGAGTGGGTGCGCAACCATTAGTAATGGTGGTAGACACACCTTC  
TATCCAGACAGTGTGAAGGGGCGATTACCATCTCCAGAGACAATGCCAAGAACACCCTGTATCTGCAAATGAGCAGT  
CTGAAGTTGAGGACACAGCCATGTATTTATGTGTAAGACAGACTGGGACGGAGGGCTGGTTTGCTTACTGGGGCCAAG  
GGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:1020)

**Mouse antibody 25E6 Heavy chain: Amino acid sequence (138 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MNFGLSLIFLALILKGVQCEVQLVESGGDLVKPGGSLKLSAASGFTFSSYGMSWVRQTPDKRLEWVATISNGGRHTF  
YPDSVKGRFTISRDNKNTLYLQMSSLKSEDTAMYLQVLRQTGTEGWFAFWGQGLVTVSA (SEQ ID NO:1021)

**Mouse antibody 25E6** heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

GAGGTGCAGCTGGTGGAGTCTGGGGGAGACTTAGTGAAGCCTGGAGGGTCCCTGAAACTCTCCTGTGCAGCCTCTGGT  
TTCACCTTTCAGT (SEQ ID NO:1022)

(amino acids)

EVQLVESGGDLVKPGGSLKLSAASGFTFS (SEQ ID NO:1023)

**Mouse antibody 25E6 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AGTTATGGAATGTCT (SEQ ID NO:1024)

(amino acids)

SYGMS (SEQ ID NO:1025)

**Mouse antibody 25E6 heavy chain variable framework 2 (FW2) sequence:**

(DNA)

TGGGTTCCGACAGACTCCAGACAAGAGGCTGGAGTGGGTCGCA (SEQ ID NO:1026)

(amino acids)

WVRQTPDKRLEWVA (SEQ ID NO:1027)

**Mouse antibody 25E6 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

ACCATTAGTAATGGTGGTAGACACACCTTCTATCCAGACAGTGTGAAGGGG (SEQ ID NO:1028)

(amino acids)

TISNGGRHTFYPSVKG (SEQ ID NO:1029)

**Mouse antibody 25E6 heavy chain variable framework 3 (FW3) sequence:**

(DNA)

CGATTACCATCTCCAGAGACAATGCCAAGAACACCCTGTATCTGCAAATGAGCAGTCTGAAGTCTGAGGACACAGCC  
ATGTATTTATGTGTAAGA (SEQ ID NO:1030)

(amino acids)

RFTISRDNKNTLYLQMSSLKSEDTAMYLVR (SEQ ID NO:1031)

**Mouse antibody 25E6 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

CAGACTGGGACGGAGGGCTGGTTTGCTTAC (SEQ ID NO:1032)

(amino acids)

QTGTEGWFAV (SEQ ID NO:1033)

**Mouse antibody 25E6 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGCCAAGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:1034)

(amino acids)

WGQGTLLTVSA (SEQ ID NO:1035)

**Mouse antibody 25E6 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAGTCCTGCCAGTTCCTGTTTCTGTTAGTGCTCTGGATTTCGGGAAACCAACGGTGATGTTGTGATGACCCAGACT  
CCACTCACTTTGTCGGTTACCATTTGACAACCAGCCTCCATCTCTTGCAAGTCAAGTCAGAGCCTCTTAGATAGTGAT  
GGAAAGACATATTTGAATTGGTTGTTACAGAGGCCAGGCCAGTCTCCAAAGCGCCTAATCTATCTGGTGTCTAAACTG  
GACTCTGGAGTCCCTGACAGGTTCACTGGCAGTGGATCAGGGACAGATTTACACTGAAAATCAGCAGAGGGAGGCTG  
AGGATTTGGGAGTTTATTATTGCTGGCAAGGTACACATTTTCTCAGACGTTTGGTGGAGGCACCAAGCTGGAAATCA  
AA (SEQ ID NO:1036)

**Mouse antibody 25E6 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MSPAQFLFLLVLWIRETNGDVVMTQTPLTSLVTIGQPASISCKSSQSLDSDGKTYLNWLLQRPQGSPKRLIYLVSKL  
DSGVPDRFTGSGSGTDFTLKISRVEAEDLGVYYCWQGTHFPQTFGGGTKLEIK (SEQ ID NO:1037)

**Mouse antibody 25E6** light chain variable *framework 1 (FW1)* sequence:

(DNA)

GATGTTGTGATGACCCAGACTCCACTCACTTTGTCGGTTACCATTTGGACAACCAGCCTCCATCTCTTGC (SEQ ID NO:1038)

(amino acids)

DVVMQTPLTSLVTIGQPASISC (SEQ ID NO:1039)

**Mouse antibody 25E6** light chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

AAGTCAAGTCAGAGCCTCTTAGATAGTGATGGAAAGACATATTTGAAT (SEQ ID NO:1040)

(amino acids)

KSSQSLDSDGKTYLN (SEQ ID NO:1041)

**Mouse antibody 25E6** light chain variable *framework 2 (FW2)* sequence:

(DNA)

TGGTTGTTACAGAGGCCAGGCCAGTCTCCAAAGCGCCTAATCTAT (SEQ ID NO:1042)

(amino acids)

WLLQRPQGSPKRLIY (SEQ ID NO:1043)

**Mouse antibody 25E6** light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

CTGGTGTCTAAACTGGACTCT (SEQ ID NO:1044)

(amino acids)

LVSKLDS (SEQ ID NO:1045)

**Mouse antibody 25E6** light chain variable *framework 3 (FW3)* sequence:

(DNA)

GGAGTCCCTGACAGGTTCACTGGCAGTGGATCAGGGACAGATTTTCACTGAAAATCAGCAGAGTGGAGGCTGAGGAT  
TTGGGAGTTTATTATTGC (SEQ ID NO:1046)

(amino acids)

GVPDRFTGSGSGTDFTLKISRVEAEDLGVYYC (SEQ ID NO:1047)

**Mouse antibody 25E6** light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

TGGCAAGGTACACATTTTCTCAGACG (SEQ ID NO:1048)

(amino acids)

WQGTHFPQT (SEQ ID NO:1049)

**Mouse antibody 25E6** light chain variable *framework 4 (FW4)* sequence:

(DNA)

TTCGGTGGAGGCACCAAGCTGGAAATCAAA (SEQ ID NO:1050)

(amino acids)

FGGGTKLEIK (SEQ ID NO:1051)

**Mouse antibody 28F9 Heavy chain: DNA sequence (399 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGGGATGGAGCTATATCATCCTCTTTTTGGTAGCAACAGCTACAGGTGTCCACTCCCAGGTCCAAGTGCAGCAGCCT  
GGGGCTGAACTGGTGCAGCCTGGGGCTTCAGTGAAGTTGTCCTGCAAGGCTTCTGGCTACACCTTCACCGGCTACTTT  
TTGTACTGGGTGAAGCAGAGGCCTGGACATGGCCTTGAGTGGATTGGGGGAATTCATCCTAGCAATGGTGATACTGAC  
TTCAATGAGAAGTTCAAGAACAAGGCCACACTGACTGTAGACATATCCTCCAGCACTGCCTACATGCAACTCAGCAGC  
CTGACATCTGAGGACTCTGCGGTCTATTATTGTACATTACTAATAGGGGTCTACTGGGGCCAAGGCACCACTCTCACA  
GTCTCCTCA (SEQ ID NO:1052)

**Mouse antibody 28F9 Heavy chain: Amino acid sequence (133 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MGWSYIIILFLVATATGVHSQVQLQQPGAELVQPGASVKLSCKASGYTFTGYFLYWVKQRPGHGLEWIGGIHPSNGDTD  
FNEKFKNKATLTVDISSSTAYMQLSSLTSEDSAVYYCTLLIGVYWGGTTLTVSS (SEQ ID NO:1053)

**Mouse antibody 28F9 heavy chain variable framework 1 (FW1) sequence:**

(DNA)

CAGGTCCAAGTGCAGCAGCCTGGGGCTGAACTGGTGCAGCCTGGGGCTTCAGTGAAGTTGTCCTGCAAGGCTTCTGGC  
TACACCTTCACC (SEQ ID NO:1054)

(amino acids)

QVQLQQPGAELVQPGASVKLSCKASGYTFT (SEQ ID NO:1055)

**Mouse antibody 28F9 heavy chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

GGCTACTTTTTGTAC (SEQ ID NO:1056)

(amino acids)

GYFLY (SEQ ID NO:1057)

**Mouse antibody 28F9 heavy chain variable framework 2 (FW2) sequence:**

(DNA)

TGGGTGAAGCAGAGGCCTGGACATGGCCTTGAGTGGATTGGG (SEQ ID NO:1058)

(amino acids)

WVKQRPGHGLEWIG (SEQ ID NO:1059)

**Mouse antibody 28F9 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

GGAATTCATCCTAGCAATGGTGATACTGACTTCAATGAGAAGTTCAAGAAC (SEQ ID NO:1060)

(amino acids)

GIHPSNGDTDFNEKFN (SEQ ID NO:1061)

**Mouse antibody 28F9 heavy chain variable framework 3 (FW3) sequence:**

(DNA)

AAGGCCACACTGACTGTAGACATATCCTCCAGCACTGCCTACATGCAACTCAGCAGCCTGACATCTGAGGACTCTGCG  
GTCTATTATTGTACATTA (SEQ ID NO:1062)

(amino acids)

KATLTVDISSSTAYMQLSSLTSEDSAVYYCTL (SEQ ID NO:1063)

**Mouse antibody 28F9 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

CTAATAGGGGTCTAC (SEQ ID NO:1064)

(amino acids)

LIGVY (SEQ ID NO:1065)

**Mouse antibody 28F9 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGCCAAGGCACCACTCTCACAGTCTCCTCA (SEQ ID NO:1066)

(amino acids)

WGQGTTLTVSS (SEQ ID NO:1067)

**Mouse antibody 28F9 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAGTCCTGCCAGTTCCTGTTTCTGTTAGTGCTCTGGATTTCGGGAAACCAACGGTGATGTTGTGATGACCCAGACT  
 CCACTCACTTTGTCGGTTACCATTGGACAACCAGCCTCCATCTCTTGCAAGTCAAGTCAGAGCCTCTTACATAGTGAT  
 GGAAAGACATATTTGATTGTTGTTTACAGAGGCCAGGCCAGTCTCCAAAGCGCCTAATCTATCTGGTGTCTAAACTG  
 GACTCTGGAGTCCCTGACAGGTTACCGGCAGTGGATCAGGGACAGATTTCACTGAAAATCAGCAGAGTGGAGGCT  
 GAGGATTTGGGAGTTTATTTTGTCTGTCAAGGTACACATTTCCGTGGACGTTCCGGTGGAGGCACCATGCTGGAAATC  
 AAA (SEQ ID NO:1068)

**Mouse antibody 28F9 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MSPAQFLFLVLWIRETNGDVMTQTPLTSLVTIGQPASISCKSSQSLLHSDGKTYLIWLLQRPQGQSPKRLIYLVSKL  
 DSGVPDRFTGSGSGTDFTLKISRVEAEDLGVYFCCQGTHFPWTFGGGTMLEIK (SEQ ID NO:1069)

**Mouse antibody 28F9 light chain variable framework 1 (FW1) sequence:**

(DNA)

GATGTTGTGATGACCCAGACTCCACTCACTTTGTCGGTTACCATTGGACAACCAGCCTCCATCTCTTGC (SEQ ID NO:1070)

(amino acids)

DVVMTQTPLTSLVTIGQPASISC (SEQ ID NO:1071)

**Mouse antibody 28F9 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AAGTCAAGTCAGAGCCTCTTACATAGTGATGGAAAGACATATTTGATT (SEQ ID NO:1072)

(amino acids)

KSSQSLLHSDGKTYLI (SEQ ID NO:1073)

**Mouse antibody 28F9 light chain variable framework 2 (FW2) sequence:**

(DNA)

TGTTGTTACAGAGGCCAGGCCAGTCTCCAAAGCGCCTAATCTAT (SEQ ID NO:1074)

(amino acids)

WLLQRPQGQSPKRLIY (SEQ ID NO:1075)

**Mouse antibody 28F9 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

CTGGTGTCTAAACTGGACTCT (SEQ ID NO:1076)

(amino acids)

LVSKLDS (SEQ ID NO:1077)

**Mouse antibody 28F9 light chain variable framework 3 (FW3) sequence:**

(DNA)

GGAGTCCCTGACAGGTTACCGGCAGTGGATCAGGGACAGATTTCACTGAAAATCAGCAGAGTGGAGGCTGAGGAT  
 TTGGGAGTTTATTTTTC (SEQ ID NO:1078)

(amino acids)

GVPDRFTGSGSGTDFTLKISRVEAEDLGVYFC (SEQ ID NO:1079)

**Mouse antibody 28F9 light chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

TGTCAGGTACACATTTCCGTGGACG (SEQ ID NO:1080)

(amino acids)



CQGTTHFPWT (SEQ ID NO:1081)

**Mouse antibody 28F9** light chain variable *framework 4 (FW4)* sequence:

(DNA)

TTCGGTGGAGGCACCATGCTGGAAATCAAA (SEQ ID NO:1082)

(amino acids)

FGGGTMLEIK (SEQ ID NO:1083)

**Mouse antibody 18B4 Heavy chain: DNA sequence (411 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGTACTTGGGACTGAACTATGTATTCATAGTTTTTCTCTTAAATGGTGTCCAGAGTGAAGTGAAACTTGAGGAGTCT  
GGAGGAGGCTTGGTGCAACCTGGGGGATCCATGAAACTCTCTTGTGCTGCCTCTGGATTCACTTTTAATGACGCCTGG  
ATGGACTGGGTCCGCCAGTCTCCAGAGAAGGGGCTTGAGTGGGTTGCTGAAATTAGAAGCACAGCTAATATTCATACA  
ACATACTATGCTGAGTCTGTCCAAGGGAGGTTCCACATCTCAAGAGATGATTCCAAAAGTAGTGTCTACCTGCAAATG  
AACAGCTTGAGAGCTGAAGACACTGGCATTATTTATTGTACCCATTACTCTACGGATTTGCTTACTGGGGCCAAGG  
ACTCTGGTCACTGTCTCTGCA (SEQ ID NO:1084)

**Mouse antibody 18B4 Heavy chain: Amino acid sequence (137 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MYLGLNYVFIVFLLNGVQSEVKLEESGGGLVQPGGSMKLSCAASGFTFNDAWMDWVRQSPEKGLEWVAEIRSTANIHT  
TYAESVQGRFTISRDDSKSSVYLQMNSLRAEDTGIYYCTPLLYGFAYWGQGLVTVSA (SEQ ID NO:1085)

**Mouse antibody 18B4** heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

GAAGTGAAACTTGAGGAGTCTGGAGGAGGCTTGGTGCAACCTGGGGGATCCATGAAACTCTCTTGTGCTGCCTCTGGA  
TTCACTTTTAAT (SEQ ID NO:1086)

(amino acids)

EVKLEESGGGLVQPGGSMKLSCAASGFTFN (SEQ ID NO:1087)

**Mouse antibody 18B4** heavy chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

GACGCCTGGATGGAC (SEQ ID NO:1088)

(amino acids)

DAWMD (SEQ ID NO:1089)

**Mouse antibody 18B4** heavy chain variable *framework 2 (FW2)* sequence:

(DNA)

TGGGTCCGCCAGTCTCCAGAGAAGGGGCTTGAGTGGGTTGCT (SEQ ID NO:1090)

(amino acids)

WVRQSPEKGLEWVA (SEQ ID NO:1091)

**Mouse antibody 18B4** heavy chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

GAAATTAGAAGCACAGCTAATATTCATACAACTACTATGCTGAGTCTGTCCAAGGG (SEQ ID NO:1092)

(amino acids)

EIRSTANIHTTYAESVQG (SEQ ID NO:1093)

**Mouse antibody 18B4** heavy chain variable *framework 3 (FW3)* sequence:

(DNA)

AGGTTCAACATCTCAAGAGATGATTCCAAAAGTAGTGTCTACCTGCAAATGAACAGCTTGAGAGCTGAAGACACTGGC  
ATTTATTATTGTACCCCA (SEQ ID NO:1094)

(amino acids)

RFTISRDDSKSSVYLQMNSLRAEDTGIYYCTP (SEQ ID NO:1095)

**Mouse antibody 18B4** heavy chain variable *complementarity determining regions 3* (CDR3) sequence:

(DNA)

TTACTCTACGGATTTGCTTAC (SEQ ID NO:1096)

(amino acids)

LLYGFAY (SEQ ID NO:1097)

**Mouse antibody 18B4** heavy chain variable *framework 4* (FW4) sequence:

(DNA)

TGGGGCCAAGGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:1098)

(amino acids)

WGQGTLVTVSA (SEQ ID NO:1099)

**Mouse antibody 18B4 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGCCTGTTAGGCTGTTGGTGCTGATGTTCTGGATTCTGCTTCCAGCAGTGATGTTGTGATGACCCAAAGT  
CCACTCTCCCTGCCTGTCTAGTCTTGGAGATCAAGCCTCCATCTCTTGCAGAACTAGTCAGAGCCTTGTACACAGTAAT  
GGAAACACCTATTTACATTGGCACCTGCAGAAGCCAGGCCAGTCTCCAAAGGTCTGATCTACAAAGTTTCCAGCCGA  
TTTTCTGGGGTCCCAGACAGGTTCACTGGCAGTGGATCGGGGACAGATTTTCACTCAAGATCAGCAGAGTGGAGGCT  
GAGGATCTGGGAGTTTATTTCTGCTCTCAAATAACACATGTTCCGTACACGTTCCGAGGGGGGACCAAGCTGGAAATA  
AAA (SEQ ID NO:1100)

**Mouse antibody 18B4 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLPVRLLLVLMFWIPASSSDVVMTQSPLSLPVSLGDAQASISCRTSQSLVHSNGNTYLHWHLQKPGQSPKVLIIYKVSSR  
FSGVPDRFSGSGSGTDFTLKISRVEAEDLGVYFCSQNTHPVPTFGGGTKLEIK (SEQ ID NO:1101)

**Mouse antibody 18B4** light chain variable *framework 1* (FW1) sequence:

(DNA)

GATGTTGTGATGACCCAAAGTCCACTCTCCCTGCCTGTCTAGTCTTGGAGATCAAGCCTCCATCTCTTGC (SEQ ID NO:1102)

(amino acids)

DVVMTQSPLSLPVSLGDAQASIS (SEQ ID NO:1103)

**Mouse antibody 18B4** light chain variable *complementarity determining regions 1* (CDR1) sequence:

(DNA)

AGAACTAGTCAGAGCCTTGTACACAGTAATGGAAACACCTATTTACAT (SEQ ID NO:1104)

(amino acids)

RTSQSLVHSNGNTYLH (SEQ ID NO:1105)

**Mouse antibody 18B4** light chain variable *framework 2* (FW2) sequence:

(DNA)

TGGCACCTGCAGAAGCCAGGCCAGTCTCCAAAGGTCTGATCTAC (SEQ ID NO:1106)

(amino acids)

WHLQKPGQSPKVLIIY (SEQ ID NO:1107)

**Mouse antibody 18B4** light chain variable *complementarity determining regions 2* (CDR2) sequence:

(DNA)

AAAGTTTCCAGCCGATTTTCT (SEQ ID NO:1108)

(amino acids)

KVSSRFS (SEQ ID NO:1109)

**Mouse antibody 18B4** light chain variable *framework 3* (FW3) sequence:

(DNA)

GGGGTCCCAGACAGGTTCACTGGCAGTGGATCGGGGACAGATTTTCACTCAAGATCAGCAGAGTGGAGGCTGAGGAT  
CTGGGAGTTTATTTCTGC (SEQ ID NO:1110)

(amino acids)  
GVPDRFSGSGSGTDFTLKISRVEAEDLGVYFC (SEQ ID NO:1111)

**Mouse antibody 18B4** light chain variable *complementarity determining regions 3* (CDR3) sequence:

(DNA)  
TCTCAAATAACACATGTTCCGTACACG (SEQ ID NO:1112)  
(amino acids)  
SQNTHVPYT (SEQ ID NO:1113)

**Mouse antibody 18B4** light chain variable *framework 4* (FW4) sequence:

(DNA)  
TTCGGAGGGGGGACCAAGCTGGAAATAAAA (SEQ ID NO:1114)  
(amino acids)  
FGGGTKLEIK (SEQ ID NO:1115)

**Mouse Antibody IE4 Heavy chain: DNA sequence (408 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4  
ATGGAATGGCCTTGTATCTTTCTCTTCTCCTGTCTAGTAAGGTGTCCACTCCCAGGTTTCAGCTGCAGCAGTCT  
GGGGCTGAGCTGGTGAGGCCTGGGTCTCAGTGAAGATTTCTGTAAAGGCTTCTGGCTATGCATTTCAGTACCTACTGG  
ATGAAGTGGGTGAAGCAGAGGCCTGGACAGGGTCTTGAGTGGATTGGACAGATTTATCTGGAGATAGTGATACTAAC  
TACAATGGAAAGTTCAAGGGTAAAGCCACACTGACTGCAGACAAGTCTCCAAACACAGCCTACATGCAGCTCAGCAGC  
CTAACATCTGAGGACTCTGCGGTCTTTTCTGTGCAAGAGGTAACCACGCCTCTATGGACTACTGGGGTCAAGGAACC  
TCAGTCACCGTCTCTCTCA (SEQ ID NO:1116)

**Mouse Antibody IE4 Heavy chain: Amino acid sequence (136 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4  
MEWPCIFLFLSVTEGVHSQVQLQQSGAELVRPGSSVKISCKASGYAFSTYWMNWVKQRPGQGLEWIGQIYPGDSDTN  
YNGKFKGKATLTADKSSNTAYMQLSSLTSEDSAVFFCARGNHASMDYWGQTSVTVSS (SEQ ID NO:1117)

**Mouse Antibody IE4** heavy chain variable *framework 1* (FW1) sequence:

(DNA)  
CAGGTTTCAGCTGCAGCAGTCTGGGGCTGAGCTGGTGAGGCCTGGGTCTCAGTGAAGATTTCTGTAAAGCCTTCTGGC  
TATGCATTTCAGT (SEQ ID NO:1118)  
(amino acids)  
QVQLQQSGAELVRPGSSVKISCKASGYAFS (SEQ ID NO:1119)

**Mouse antibody IE4** heavy chain variable *complementarity determining regions 1* (CDR1) sequence:

(DNA)  
ACCTACTGGATGAAC (SEQ ID NO:1200)  
(amino acids)  
TYWMN (SEQ ID NO:1201)

**Mouse antibody IE4** heavy chain variable *framework 2* (FW2) sequence:

(DNA)  
TGGGTGAAGCAGAGGCCTGGACAGGGTCTTGAGTGGATTGGA (SEQ ID NO:1202)  
(amino acids)  
WVKQRPGQGLEWIG (SEQ ID NO:1203)

**Mouse antibody IE4** heavy chain variable *complementarity determining regions 2* (CDR2) sequence:

(DNA)  
CAGATTTATCTGGAGATAGTGATACTAACTACAATGGAAAGTTCAAGGGT (SEQ ID NO:1204)  
(amino acids)  
QIYPGDSDTNYNGKFKG (SEQ ID NO:1205)

**Mouse antibody IE4 heavy chain variable framework 3 (FW3) sequence:**

(DNA)

AAAGCCCACTGACTGCAGACAAGTCCTCCAACACAGCCTACATGCAGCTCAGCAGCCTAACATCTGAGGACTCTGCG  
GTCTTTTCTGTGCAAGA (SEQ ID NO:1206)

(amino acids)

KATLTADKSSNTAYMQLSSLTSEDSAVFFCAR (SEQ ID NO:1207)

**Mouse antibody IE4 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

GGTAACCACGCCTCTATGGACTAC (SEQ ID NO:1208)

(amino acids)

GNHASMDY (SEQ ID NO:1209)

**Mouse antibody IE4 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGTCAAGGAACCTCAGTCACCGTCTCCTCA (SEQ ID NO:1210)

(amino acids)

WGQGTSVTVSS (SEQ ID NO:1211)

**Mouse Antibody IE4 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGCCTGTTAGGCTGTTGGTGCTGATGTTCTGGATTCTGCTTCCAGCAGTGATGTTGTGATGACCCAACT  
CCACTCTCCCTGCCTGTCAGTCTTGGAGATCAAGCCTCCATCTCTTGCAGATCTAGTCAGAGCCTTGTACACAGTAAT  
GGAAACACCTATTTACATTGGTACCTGCAGAAGCCAGGCCAGTCTCCAAAGCTCCTGATCTACAAAGTTTCCAACCGA  
TTTTCTGGGGTCCCAGACAGGTTCAAGTGGCAGTGGATCAGGGACAGATTTCACTCAAGATCAGCAGAGTGGAGGCT  
GAGGATCTGGGAGTTTATTTCTGCTCTCAAAAAACACATGTTCCGTGGACGTTCCGGTGGAGGCACCAAGCTGGAAATC  
AAA (SEQ ID NO:1212)**Mouse Antibody IE4 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLPVRLLVLMFWIPASSSDVMTQTPLSLPVSLGDAQASISCRSSQSLVHSNGNTYLHWYLQKPGQSPKLLIYKVSNR  
FSGVPDRFSGSGSGTDFTLKISRVEAEDLGVYFCSQKTHVPWTFGGGTKLEIK (SEQ ID NO:1213)**Mouse antibody IE4 light chain variable framework 1 (FW1) sequence:**

(DNA)

GATGTTGTGATGACCCAACTCCACTCTCCCTGCCTGTCAGTCTTGGAGATCAAGCCTCCATCTCTTGC (SEQ ID  
NO:1214)

(amino acids)

DVVMTQTPLSLPVSLGDAQASISC (SEQ ID NO:1215)

**Mouse antibody IE4 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AGATCTAGTCAGAGCCTTGTACACAGTAATGGAAACACCTATTTACAT (SEQ ID NO:1216)

(amino acids)

RSSQSLVHSNGNTYLH (SEQ ID NO:1217)

**Mouse antibody IE4 light chain variable framework 2 (FW2) sequence:**

(DNA)

TGGTACCTGCAGAAGCCAGGCCAGTCTCCAAAGCTCCTGATCTAC (SEQ ID NO:1218)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:1219)

**Mouse antibody IE4 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

AAAGTTTCCAACCGATTTTCT (SEQ ID NO:1220)  
(amino acids)  
KVSNRFS (SEQ ID NO:1221)

**Mouse antibody IE4** light chain variable *framework 3 (FW3)* sequence:

(DNA)  
GGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAGGGACAGATTTTCACTCAAGATCAGCAGAGTGGAGGCTGAGGAT  
CTGGGAGTTTATTTCTGC (SEQ ID NO:1222)  
(amino acids)  
GVPDRFSGSGSGTDFTLKISRVEAEDLGVYFC (SEQ ID NO:1223)

**Mouse antibody IE4** light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)  
TCTCAAAAAACACATGTTCCGTGGACG (SEQ ID NO:1224)  
(amino acids)  
SQKTHVPWT (SEQ ID NO:1225)

**Mouse antibody IE4** light chain variable *framework 4 (FW4)* sequence:

(DNA)  
TTCGGTGGAGGCACCAAGCTGGAAATCAAA (SEQ ID NO:1226)  
(amino acids)  
FGGGTKLEIK (SEQ ID NO:1227)

**Mouse antibody 29H1 Heavy chain: DNA sequence (411 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4  
ATGTACTTGGGACTGAACTATGTATTCATAGTTTTTCTCTTAAATGGTGTCCAGAGTGAAGTGAAGCTTGAGGAGTCT  
GGAGGAGGCTTGGTACAACCTGGAGGATCCATGAACTCTCTTGTGCTGCCTCTGGATTCACTTTTAGTGACGCCTGG  
ATGGACTGGGTCCGCCAGTCTCCAGAGAAGGGGCTTGAATGGGTTGCTGAAATTAGAAGCAAAGCTACTAATCATGCA  
ACATACTATGCTGAGTCTGTGAAAGGGAGGTTCCACCATCTCAAGAGATGATTCCAAAAGTAGTGTCTACCTGCAAATG  
AACAGCTTAAGAGCTGAAGACACTGGCATTATTTACTGTACCCCCCTACTTTACGGGTTTGCTTACTGGGGCCAAGGG  
ACTCTGGTCACTGTCTCTGCA (SEQ ID NO:1228)

**Mouse antibody 29H1 Heavy chain: Amino acid sequence (137 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4  
MYLGLNYVFIVFLLNGVQSEVKLEESGGGLVQPGGSMKLSCAASGFTFSDAWMDWVRQSPKGLEWVAEIRSKATNHA  
TYAESVKGRFTISRDDSKSSVYLQMNSLRAEDTGIYYCTPLLYGFAYWGQGLVTVSA (SEQ ID NO:1229)

**Mouse antibody 29H1** heavy chain variable *framework 1 (FW1)* sequence:

(DNA)  
GAAGTGAAGCTTGAGGAGTCTGGAGGAGGCTTGGTACAACCTGGAGGATCCATGAACTCTCTTGTGCTGCCTCTGGA  
TTCACCTTTTAGT (SEQ ID NO:1230)  
(amino acids)  
EVKLEESGGGLVQPGGSMKLSCAASGFTFS (SEQ ID NO:1231)

**Mouse antibody 29H1** heavy chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)  
GACGCCTGGATGGAC (SEQ ID NO:1232)  
(amino acids)  
DAWMD (SEQ ID NO:1233)

**Mouse antibody 29H1** heavy chain variable *framework 2 (FW2)* sequence:

(DNA)  
TGGGTCCGCCAGTCTCCAGAGAAGGGGCTTGAATGGGTTGCT (SEQ ID NO:1234)  
(amino acids)

WVRQSPEKGLEWVA (SEQ ID NO:1235)

**Mouse antibody 29H1 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

GAAATTAGAAGCAAAGCTACTAATCATGCAACATACTATGCTGAGTCTGTGAAAGGG (SEQ ID NO:1236)

(amino acids)

EIRSKATNHATYYAESVKG (SEQ ID NO:1237)

**Mouse antibody 29H1 heavy chain variable framework 3 (FW3) sequence:**

(DNA)

AGGTTCCACCATCTCAAGAGATGATTCCAAAAGTAGTGTCTACCTGCAAATGAACAGCTTAAGAGCTGAAGACACTGGC

ATTATTACTGTACCCCC (SEQ ID NO:1238)

(amino acids)

RFTISRDDSKSSVYLQMNSLRAEDTGIYYCTP (SEQ ID NO:1239)

**Mouse antibody 29H1 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

CTACTTTACGGGTTTGCTTAC (SEQ ID NO:1240)

(amino acids)

LLYGFAY (SEQ ID NO:1241)

**Mouse antibody 29H1 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGCCAAGGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:1242)

(amino acids)

WGQGTILVTVSA (SEQ ID NO:1243)

**Mouse antibody 29H1 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGCCTGTTAGGCTGTTGGTGCTGATGTTCTGGATTCTGCTTCCAGCAGTGATGTTGTGATGACCCAAACT  
CCACTCTCCCTGCCTGTCAGTCTTGGAGATCAAGCCTCCATCTCTTGCAGATCTGGTCAGAGCCTTGACACAGTAAT  
GGACACACCTATTTACATTGGTACCTGCAGAAGCCAGGCCAGTCTCCAAGGCTCCTGATCTACAAAGTTTCCAACCGA  
TTTTCTGGGGTCCCAGACAGGTTCACTGGCAGTGGATCAAGGGCAGATTTTCACTCAAGATCAGCAGAGTGGAGGCT  
GAGGATCTGGGAGTTTATTTCTGCTCTCAAACCTACACATGTTCCGTGGACGTTCCGGTGGAGGCACCAAGCTGGAAATC  
AAA (SEQ ID NO:1244)

**Mouse antibody 29H1 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLPVRLLLVLMFWIPASSSDVVMQTPLSLPVSLGDAQASISCRSGQSLVHSNGHTYLNHWYLNKPGQSPRLLIYKVSNR  
FSGVPDRFSGSGSRADFTLKISRVEAEDLGVYFCSQTHVPWTFGGGTKLEIK (SEQ ID NO:1245)

**Mouse antibody 29H1 light chain variable framework 1 (FW1) sequence:**

(DNA)

GATGTTGTGATGACCCAACTCCACTCTCCCTGCCTGTCAGTCTTGGAGATCAAGCCTCCATCTCTTGC (SEQ ID NO:1246)

(amino acids)

DVVMQTPLSLPVSLGDAQASIS (SEQ ID NO:1247)

**Mouse antibody 29H1 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AGATCTGGTCAGAGCCTTGTACACAGTAATGGACACACCTATTTACAT (SEQ ID NO:1248)

(amino acids)

RSGQSLVHSNGHTYLH (SEQ ID NO:1249)

**Mouse antibody 29H1** light chain variable *framework 2 (FW2)* sequence:

(DNA)

TGGTACCTGCAGAAGCCAGGCCAGTCTCCAAGGCTCCTGATCTAC (SEQ ID NO:1250)

(amino acids)

WYLQKPGQSPRLLIY (SEQ ID NO:1251)

**Mouse antibody 29H1** light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

AAAGTTTCCAACCGATTTTCT (SEQ ID NO:1252)

(amino acids)

KVSNRFS (SEQ ID NO:1253)

**Mouse antibody 29H1** light chain variable *framework 3 (FW3)* sequence:

(DNA)

GGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAAGGGCAGATTTCACTCAAGATCAGCAGAGTGGAGGCTGAGGATCTGGGAGTTTATTTCTGC (SEQ ID NO:1254)

(amino acids)

GVPDRFSGSGSRADFTLKISRVEAEDLGVYFC (SEQ ID NO:1255)

**Mouse antibody 29H1** light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

TCTCAAACCTACACATGTTCCGTGGACG (SEQ ID NO:1256)

(amino acids)

SQTTHVPWT (SEQ ID NO:1257)

**Mouse antibody 29H1** light chain variable *framework 4 (FW4)* sequence:

(DNA)

TTCGGTGGAGGCACCAAGCTGGAAATCAAA (SEQ ID NO:1258)

(amino acids)

FGGGTKLEIK (SEQ ID NO:1259)

**Mouse antibody 31A1 Heavy chain: DNA sequence (399 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGGAAAGGCACTGGATCTTTCTCTTCCTGTTTTTCAGTAAGTGCAGGTGTCCACTCCCAGGTCCAGCTTCAGCAGTCTGGGGCTGAACTGGCAAAACCTGGGGCCTCAGTGAAGATGTCCTGCAAGGCTTCTGGCTACACCTTTACTAGCTACTGGATGCACTGGGTAAAACAGAGGCCTGGACAGGGTCTGGAATGGATTGGATACATTAATCCTAGCACTGGTTATACTGAGTACAATCAGAAGTTCAAGGACAAGGCCACATTGACTGCAGACAAATCCTCCAGCACAGCCTACATGCAACTGAGCAGCCTGACATCTGAGGACTCTGCAGTCTATTACTGTGCAAGAGCCTACATTGACTACTGGGGCCAAGGCACCACTCTCACA GTCTCCTCA (SEQ ID NO:1260)

**Mouse antibody 31A1 Heavy chain: Amino acid sequence (133 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MERHWIFLFLFSVTAGVHSQVQLQQSGAELAKPGASVKMSCKASGYTFTSYWMHWVKQRPGQGLEWIGYINPSTGYTE  
YNQKFKDKATLTADKSSSTAYMQLSSLTSEDSAVYYCARAYIDYWGQGTTTLTVSS (SEQ ID NO:1261)

**Mouse antibody 31A1** heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

CAGGTCCAGCTTCAGCAGTCTGGGGCTGAACTGGCAAAACCTGGGGCCTCAGTGAAGATGTCTTGCAAGGCTTCTGGC  
TACACCTTTACT (SEQ ID NO:1262)

(amino acids)

QVQLQQSGAELAKPGASVKMSCKASGYTFT (SEQ ID NO:1263)

**Mouse antibody 31A1** heavy chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

AGCTACTGGATGCAC (SEQ ID NO:1264)

(amino acids)

SYWMH (SEQ ID NO:1265)

**Mouse antibody 31A1** heavy chain variable *framework 2 (FW2)* sequence:

(DNA)

TGGGTAAAACAGAGGCCTGGACAGGGTCTGGAATGGATTGGA (SEQ ID NO:1266)

(amino acids)

WVKQRPGQGLEWIG (SEQ ID NO:1267)

**Mouse antibody 31A1** heavy chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

TACATTAATCCTAGCACTGGTTATACTGAGTACAATCAGAAGTTCAAGGAC (SEQ ID NO:1268)

(amino acids)

YINPSTGYTEYNQKFKD (SEQ ID NO:1269)

**Mouse antibody 31A1** heavy chain variable *framework 3 (FW3)* sequence:

(DNA)

AAGGCCACATTGACTGCAGACAAATCCTCCAGCACAGCCTACATGCAACTGAGCAGCCTGACATCTGAGGACTCTGCA  
GTCTATTACTGTGCAAGA (SEQ ID NO:1270)

(amino acids)

KATLTADKSSSTAYMQLSSLTSEDSAVYYCAR (SEQ ID NO:1271)

**Mouse antibody 31A1** heavy chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

GCCTACATTGACTAC (SEQ ID NO:1272)

(amino acids)

AYIDY (SEQ ID NO:1273)

**Mouse antibody 31A1** heavy chain variable *framework 4 (FW4)* sequence:

(DNA)

TGGGGCCAAGGCACCACTCTCACAGTCTCCTCA (SEQ ID NO:1274)

(amino acids)



WGQGTTLTVSS (SEQ ID NO:1275)

**Mouse antibody 31A1 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGCCTGTTAGGCTGTTGGTGCTGATGTTCTGGATTCTGCTTCCAGCAGTGATGTTTTGATGACCCAAACT  
CCACTCTCCCTGCCTGTCAGTCTTGGAGATCAAGCCTCCTTCTCTTGCAGATCTAGTCAGAGCATTGTACATAGTAAT  
GGAAACACCTATTTAGAATGGTACCTGCAGAAACCAGGCCAGTCTCCAAAGCTCCTGATCTACAAAGTTTCCAACCGA  
TTTTCTGGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAGGGACAGATTTTCACTCAAGATCAACAGAGTGGAGGCT  
GAGGATCTGGGAGTTTATTACTGCTTTCAAGTTTACATTTTCCGTGGACGTTCCGGTGGAGGCACCAAGCTGGAAATC  
AAA (SEQ ID NO:1276)

**Mouse antibody 31A1 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLPVRLLLVLMFWIPASSSDVLMQTPLSLPVSLGDAQSFSCRSSQSIVHSNGNTYLEWYLQKPGQSPKLLIYKVSNR  
FSGVPDRFSGSGSGTDFTLKINRVEAEDLGVIYCFQVSHFPWTFGGGTKLEIK (SEQ ID NO:1277)

**Mouse antibody 31A1 light chain variable framework 1 (FW1) sequence:**

(DNA)

GATGTTTTGATGACCCAAACTCCACTCTCCCTGCCTGTCAGTCTTGGAGATCAAGCCTCCTTCTCTTGC (SEQ ID NO:1278)

(amino acids)

DVLMQTPLSLPVSLGDAQSFSC (SEQ ID NO:1279)

**Mouse antibody 31A1 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AGATCTAGTCAGAGCATTGTACATAGAAATGGAAACACCTATTTAGAA (SEQ ID NO:1280)

(amino acids)

RSSQSIVHRNGNTYLE (SEQ ID NO:1281)

**Mouse antibody 31A1 light chain variable framework 2 (FW2) sequence:**

(DNA)

TGGTACCTGCAGAAACCAGGCCAGTCTCCAAAGCTCCTGATCTAC (SEQ ID NO:1282)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:1283)

**Mouse antibody 31A1 light chain variable complementarity determining regions 2 (CDR2) sequence:**

(DNA)

AAAGTTTCCAACCGATTTTCT (SEQ ID NO:1284)

(amino acids)

KVSNRFS (SEQ ID NO:1285)

**Mouse antibody 31A1 light chain variable framework 3 (FW3) sequence:**

(DNA)

GGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAGGGACAGATTTTCACTCAAGATCAACAGAGTGGAGGCTGAGGAT  
CTGGGAGTTTATTACTGC (SEQ ID NO:1286)

(amino acids)

GVPDRFSGSGSGTDFTLKINRVEAEDLGVIYCFQVSHFPWTFGGGTKLEIK (SEQ ID NO:1287)

**Mouse antibody 31A1** light chain variable *complementarity determining regions 3* (CDR3) sequence:

(DNA)

TTTCAAGTTTCACATTTTCCGTGGACG (SEQ ID NO:1288)

(amino acids)

FQVSHFPWT (SEQ ID NO:1289)

**Mouse antibody 31A1** light chain variable *framework 4* (FW4) sequence:

(DNA)

TTCGGTGGAGGCACCAAGCTGGAAATCAAA (SEQ ID NO:1290)

(amino acids)

FGGGTKLEIK (SEQ ID NO:1291)

**Mouse antibody 32C1 Heavy chain: DNA sequence (411 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGTACTTGGGACTGAACTGTGTATTCATAGTTTTTCTCTTAAAGGTGTCCAGAGTGAAGTGAAGCTTGAGGAGTCT  
GGAGGAGGCTTGGTGCAATCTGGAGGATCCATGAACTCTCCTGTGTGCTTGGCTCTGGATTCACTTTCAGTAATTACTGG  
ATGAACTGGGTCCGCCAGTCTCCAGAGAAGGGGCTTGAGTGGGTTGCTGAAATTAGATTGAAATCTAATAATTATGCA  
ATACATTATGCGGAGTCTGTGAAGGGGAGGTTACCATCTCAAGAGATGATTCCAAAAGTAGTGTCTACCTGCAAATG  
AACAACCTTAAGAGCTGAAGACACTGGCATTATTTACTGTACCAGGGTCCCGGGACTGGATGCTTACTGGGGCCAAGG  
ACTCTGGTCACTGTCTCTGCA (SEQ ID NO:1292)

**Mouse antibody 32C1 Heavy chain: Amino acid sequence (137 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MYLGLNCVFIVFLLKGVQSEVKLEESGGGLVQSGGSMKLSCVASGFTFSNYWMNWVRQSPEKGLEWVAEIRLKSNNYA  
IHVAESVKGRFTISRDDSKSSVYLQMNNLRAEDTGIYYCTRVPLDAYWGQGLVTVSA (SEQ ID NO:1293)

**Mouse antibody 32C1** heavy chain variable *framework 1* (FW1) sequence:

(DNA)

GAAGTGAAGCTTGAGGAGTCTGGAGGAGGCTTGGTGCAATCTGGAGGATCCATGAACTCTCCTGTGTGCTTGCCTCTGGA  
TTCACCTTTCAGT (SEQ ID NO:1294)

(amino acids)

EVKLEESGGGLVQSGGSMKLSCVASGFTFS (SEQ ID NO:1295)

**Mouse antibody 32C1** heavy chain variable *complementarity determining regions 1* (CDR1) sequence:

(DNA)

AATTACTGGATGAAC (SEQ ID NO:1296)

(amino acids)

NYWMN (SEQ ID NO:1297)

**Mouse antibody 32C1** heavy chain variable *framework 2* (FW2) sequence:

(DNA)

TGGGTCCGCCAGTCTCCAGAGAAGGGGCTTGAGTGGGTTGCT (SEQ ID NO:1298)

(amino acids)

WVRQSPEKGLEWVA (SEQ ID NO:1299)

**Mouse antibody 32C1** heavy chain variable *complementarity determining regions 2* (CDR2) sequence:

(DNA)

GAAATTAGATTGAAATCTAATAATTATGCAATACATTATGCGGAGTCTGTGAAGGGG (SEQ ID NO:1300)

(amino acids)

EIRLKSNNYAIHYAESVKG (SEQ ID NO:1301)

**Mouse antibody 32C1 heavy chain variable framework 3 (FW3) sequence:**

(DNA)

AGGTTCCACCATCTCAAGAGATGATTCCAAAAGTAGTGTCTACCTGCAAATGAACAACTTAAGAGCTGAAGACACTGGC  
ATTATTACTGTACCAGG (SEQ ID NO:1302)

(amino acids)

RFTISRDDSKSSVYLQMNRLRAEDTGIYYCTR (SEQ ID NO:1303)

**Mouse antibody 32C1 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**

(DNA)

GTCCCGGGACTGGATGCTTAC (SEQ ID NO:1304)

(amino acids)

VPGLDAY (SEQ ID NO:1305)

**Mouse antibody 32C1 heavy chain variable framework 4 (FW4) sequence:**

(DNA)

TGGGGCCAAGGGACTCTGGTCACTGTCTCTGCA (SEQ ID NO:1306)

(amino acids)

WGQGTILVTVSA (SEQ ID NO:1307)

**Mouse antibody 32C1 Light chain: DNA sequence (393 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAGTTGCCTGTTAGGCTGTTGGTGCTGATGTTCTGGATTCCTTCCAGCAGTGATGTTGTGATGACCCAAACT  
CCACTCTCCCTGCCTGTCAGTCTTGGAGATCAAGCCTCCATCTCTTGCAGATCTAGTCAGAGCCTTGTACACAGTAAT  
GGAAACACCTATTTACATTGGTACCTGCAGAAGCCAGGCCAGTCTCCAAAGCTCCTGATCTACAAAGTTTCCAACCGA  
TTTTCTGGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAGGGACAGATTTTCACTCAAGATCAGCAGTGTGGAGGCT  
GAGGATCTGGGAGTTTATTTCTGCTCTCAAATTACACATGTTCCGTACACGTTCCGAGGGGGGACCAATCTGGAAATA  
AAA (SEQ ID NO:1308)

**Mouse antibody 32C1 Light chain: Amino acid sequence (131 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKLPVRLLVLMFWIPASSSDVVMQTPLSLPVSLGDAQASISCRSSQSLVHSNGNTYLHWYLQKPGQSPKLLIYKVSNR  
FSGVPDRFSGSGSGTDFTLKISSVEAEDLGVYFCSQITHVPYTFGGGTNLEIK (SEQ ID NO:1309)

**Mouse antibody 32C1 light chain variable framework 1 (FW1) sequence:**

(DNA)

GATGTTGTGATGACCCAACTCCACTCTCCCTGCCTGTCAGTCTTGGAGATCAAGCCTCCATCTCTTGC (SEQ ID  
NO:1310)

(amino acids)

DVVMQTPLSLPVSLGDAQASISC (SEQ ID NO:1311)

**Mouse antibody 32C1 light chain variable complementarity determining regions 1 (CDR1) sequence:**

(DNA)

AGATCTAGTCAGAGCCTTGTACACAGTAATGGAAACACCTATTTACAT (SEQ ID NO:1312)

(amino acids)

RSSQSLVHSNGNTYLH (SEQ ID NO:1313)

**Mouse antibody 32C1 light chain variable framework 2 (FW2) sequence:**

(DNA)

TGGTACCTGCAGAAGCCAGGCCAGTCTCCAAAGCTCCTGATCTAC (SEQ ID NO:1314)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:1315)

**Mouse antibody 32C1** light chain variable *complementarity determining regions 2* (CDR2) sequence:

(DNA)

AAAGTTTCCAACCGATTTTCT (SEQ ID NO:1316)

(amino acids)

KVSNRFS (SEQ ID NO:1317)

**Mouse antibody 32C1** light chain variable *framework 3* (FW3) sequence:

(DNA)

GGGGTCCCAGACAGGTTTCAGTGGCAGTGGATCAGGGACAGATTTACACTCAAGATCAGCAGTGTGGAGGCTGAGGATCTGGGAGTTTATTTCTGC (SEQ ID NO:1318)

(amino acids)

GVPDRFSGSGSGTDFTLKISSVEAEDLGVYFC (SEQ ID NO:1319)

**Mouse antibody 32C1** light chain variable *complementarity determining regions 3* (CDR3) sequence:

(DNA)

TCTCAAATTACACATGTTCCGTACACG (SEQ ID NO:1320)

(amino acids)

SQITHVPYT (SEQ ID NO:1321)

**Mouse antibody 32C1** light chain variable *framework 4* (FW4) sequence:

(DNA)

TTCGGAGGGGGACCAATCTGGAAATAAAA (SEQ ID NO:1322)

(amino acids)

FGGGTNLEIK (SEQ ID NO:1323)

**Mouse antibody 45C11 Heavy chain: DNA sequence (423 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

ATGAAATGCAGCTGGGTTATCTTCTCCTGATGGCAGTGGTTACAGGGGTCAATTCAGAGGTTTCAGCTGCAGCAGTCTGGGGCAGACCTTGTGAAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTTTATGCACTGGGTGAAGCAGAGGCCTGAACAGGGCCTGGAGTGGATTGGAAGGATTGATCCTGCGAATGGTAATACTAAATATGACCCGAAATTCCAGGGCAAGGCCACTATAACAGCAGACACATCCTCCAACACAGCCTACCTGCAGCTCAGCAGCCTGACATCTGAGGACACTGCCGTCTATTACTGTGCTAAACCGTATGGTAACTACGGCTATTACTATGCTTTGGACTACTGGGGTCAAGGAACCTCAGTCACCGTCTCCTCA (SEQ ID NO:1324)

**Mouse antibody 45C11 Heavy chain: Amino acid sequence (141 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4

MKCSWVIFFLMAVVTGVNSEVQLQQSGADLVKPGASVKLSCTASGFNIKDTFMHWVKQRPEQGLEWIGRIDPANGNTKYDPKFQ GKATITADTSSNTAYLQLSSLTSEDTA VYYCAKPYGNYGYYYALDYWGQGTSTVTVSS (SEQ ID NO:1325)

**Mouse antibody 45C11** heavy chain variable *framework 1* (FW1) sequence:

(DNA)

GAGGTTTCAGCTGCAGCAGTCTGGGGCAGACCTTGTGAAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAA (SEQ ID NO:1326)

(amino acids)

EVQLQQSGADLVKPGASVKLSCTASGFNIK (SEQ ID NO:1327)

**Mouse antibody 45C11** heavy chain variable *complementarity determining regions 1* (CDR1) sequence:

(DNA)

GACACCTTTATGCAC (SEQ ID NO:1328)  
 (amino acids)  
 DTFMH (SEQ ID NO:1329)

**Mouse antibody 45C11 heavy chain variable framework 2 (FW2) sequence:**  
 (DNA)  
 TGGGTGAAGCAGAGGCCTGAACAGGGCCTGGAGTGGATTGGA (SEQ ID NO:1330)  
 (amino acids)  
 WVKQRPEQGLEWIG (SEQ ID NO:1331)

**Mouse antibody 45C11 heavy chain variable complementarity determining regions 2 (CDR2) sequence:**  
 (DNA)  
 AGGATTGATCCTGCGAATGGTAATACTAAATATGACCCGAAATTCCAGGGC (SEQ ID NO:1332)  
 (amino acids)  
 RIDPANGNTKYDPKFQG (SEQ ID NO:1333)

**Mouse antibody 45C11 heavy chain variable framework 3 (FW3) sequence:**  
 (DNA)  
 AAGGCCACTATAACAGCAGACACATCCTCCAACACAGCCTACCTGCAGCTCAGCAGCCTGACATCTGAGGACACTGCC  
 GTCTATTACTGTGCTAAA (SEQ ID NO:1334)  
 (amino acids)  
 KATITADTSSNTAYLQLSSLTSEDYAVYYCAK (SEQ ID NO:1335)

**Mouse antibody 45C11 heavy chain variable complementarity determining regions 3 (CDR3) sequence:**  
 (DNA)  
 CCGTATGGTAACTACGGCTATTACTATGCTTTGGACTAC (SEQ ID NO:1336)  
 (amino acids)  
 PYGNYGYYYALDY (SEQ ID NO:1337)

**Mouse antibody 45C11 heavy chain variable framework 4 (FW4) sequence:**  
 (DNA)  
 TGGGGTCAAGGAACCTCAGTCACCGTCTCCTCA (SEQ ID NO:1338)  
 (amino acids)  
 WGQGTSTVTVSS (SEQ ID NO:1339)

**Mouse antibody 45C11 Light chain: DNA sequence (381 bp)**

Signal sequence-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4  
 ATGAGGTTCCAGGTTTCAGGTTCTGGGGCTCCTTCTGCTCTGGATATCAGGTGCCCAGTGTGATGTCCAGATAACCCAG  
 TCTCCATCTTATCTTGCTGCATCTCCTGGAGAAACCATTACTATTAATTGCAGGGCAAGTAAGAGCATTAGCAAATAT  
 TTAGCCTGGTATCAAGAGAAACCTGGGAAAATAAAGCTTCTTATCTACTCTGGATCCACTTTGCAATCTGGAATT  
 CCATCAAGGTTTCAGTGGCAGTGGATCTGGTACAGATTTCACTCTCACCATCAGTAGCCTGGAGCCTGAAGATTTTGCA  
 ATGTATTACTGTCAACAGCATAATGAATTCCTGGACGTTCTGGTGGAGGCACCAAGCTGGAAATCAAA (SEQ ID  
 NO:1340)

**Mouse antibody 45C11 Light chain: Amino acid sequence (127 aa)**

Signal peptide-FR1-CDR1-FR2-CDR2-FR3-CDR3-FR4  
 MRFQVQVLGLLLLWISGAQCDVQITQSPSYLAASPGETITINCRASKSISKYLAWYQEKPGKTNKLLIYSGSTLQSGI  
 PSRFSGSGSGTDFLTITSSLEPEDFAMYYCQQHNEFPWTFGGGTKLEIK (SEQ ID NO:1341)

**Mouse antibody 45C11 light chain variable framework 1 (FW1) sequence:**

(DNA)  
 GATGTCCAGATAACCCAGTCTCCATCTTATCTTGCTGCATCTCCTGGAGAAACCATTACTATTAATTGC (SEQ ID  
 NO:1342)  
 (amino acids)  
 DVQITQSPSYLAASPGETITINC (SEQ ID NO:1343)

**Mouse antibody 45C11** light chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

AGGGCAAGTAAGAGCATTAGCAAATATTTAGCC (SEQ ID NO:1344)

(amino acids)

RASKSISKYLA (SEQ ID NO:1345)

**Mouse antibody 45C11** light chain variable *framework 2 (FW2)* sequence:

(DNA)

TGGTATCAAGAGAAACCTGGGAAACTAATAAGCTTCTTATCTAC (SEQ ID NO:1346)

(amino acids)

WYQEKPGKTNKLLIY (SEQ ID NO:1347)

**Mouse antibody 45C11** light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

TCTGGATCCACTTTGCAATCT (SEQ ID NO:1348)

(amino acids)

SGSTLQS (SEQ ID NO:1349)

**Mouse antibody 45C11** light chain variable *framework 3 (FW3)* sequence:

(DNA)

GGAATTCCATCAAGGTTTCAGTGGCAGTGGATCTGGTACAGATTTCACTCTCACCATCAGTAGCCTGGAGCCTGAAGAT

TTTGCAATGTATTACTGT (SEQ ID NO:1350)

(amino acids)

GIPSRFSGSGSGTDFLTITSSLEPEDFAMYYC (SEQ ID NO:1351)

**Mouse antibody 45C11** light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

CAACAGCATAATGAATTCCCGTGGACG (SEQ ID NO:1352)

(amino acids)

QQHNEFPWT (SEQ ID NO:1353)

**Mouse antibody 45C11** light chain variable *framework 4 (FW4)* sequence:

(DNA)

TTCGGTGGAGGCACCAAGCTGGAAATCAAA (SEQ ID NO:1354)

(amino acids)

FGGGTKLEIK (SEQ ID NO:1355)

5C6F3

Mouse 5C6F3 heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

Gaagtgatgctggtggagtcctgggggaggetttagtgaagcctggaggggtccctgaaactctcctgtgcagcctctgga

ttcactttcagt (SEQ ID NO:1356)

(amino acids)

EVMLVESGGGLVKPGGSLKLSAASGFTFS (SEQ ID NO:1357)

Mouse 5C6F3 heavy chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

acctatgccatgtct (SEQ ID NO:1358)

(amino acids)

TYAMS (SEQ ID NO:1359)

Mouse 5C6F3 heavy chain variable *framework 2 (FW2)* sequence:

(DNA)

tgggttcgccagactccggagaagaggctggagtgggtcgca (SEQ ID NO:1360)  
(amino acids)  
WVRQTPEKRLEWVA (SEQ ID NO:1361)

Mouse 5C6F3 heavy chain variable *complementarity determining regions 2 (CDR2)*  
sequence:  
(DNA)  
gccattagtaatggtggtggttacacctactatccagacagtctgaagggg (SEQ ID NO:1362)  
(amino acids)  
AISNGGGYTYYPDSLKG (SEQ ID NO:1363)

Mouse 5C6F3 heavy chain variable *framework 3 (FW3)* sequence:  
(DNA)  
cgattcaccatctccagagacaatgccaaagacacctgtacctgcaaagagcagtcctgaggtctgaggacacggcc  
acgtattactgtgcaaga (SEQ ID NO:1364)  
(amino acids)  
RFTISRDNAKNTLYLQMSSLRSEDTATYYCAR (SEQ ID NO:1365)

Mouse 5C6F3 heavy chain variable *complementarity determining regions 3 (CDR3)*  
sequence:  
(DNA)  
cgttactatgatcactactttgactac (SEQ ID NO:1366)  
(amino acids)  
RYYDHYFDY (SEQ ID NO:1367)

Mouse 5C6F3 heavy chain variable *framework 4 (FW4)* sequence:  
(DNA)  
tggggccaaggcaccgctctcacggtctctctca (SEQ ID NO:1368)  
(amino acids)  
WGQGTALTIVSS (SEQ ID NO:1369)

Mouse 5C6F3 light chain variable *framework 1 (FR1)* sequence:  
(DNA)  
gatgttttgatgacccaaactccactctccctgcctgtcagtccttgagatcaagcctccatctcttgc (SEQ ID  
NO:1370)  
(amino acids)  
DVLMTQTPLSLPVSLGDQASISC (SEQ ID NO:1371)

Mouse 5C6F3 light chain variable *complementarity determining regions 1 (CDR1)*  
sequence:  
(DNA)  
agatctagtcagaccattgtacatagtaatggaaacacctatthagaa (SEQ ID NO:1372)  
(amino acids)  
RSSQTIVHSNGNTYLE (SEQ ID NO:1373)

Mouse 5C6F3 light chain variable *framework 2 (FR2)* sequence:  
(DNA)  
tggtacctgcagaaaccaggccagtcctccaaagctcctgatctac (SEQ ID NO:1374)  
(amino acids)  
WYLQKPGQSPKLLIY (SEQ ID NO:1375)

Mouse 5C6F3 light chain variable *complementarity determining regions 2 (CDR2)*  
sequence:  
(DNA)  
aaagtttccaaccgattttct (SEQ ID NO:1376)  
(amino acids)  
KVSNRFS (SEQ ID NO:1377)

Mouse 5C6F3 light chain variable *framework 3 (FR3)* sequence:

(DNA)

ggggtcccagacaggttcagtgccagtgatcagggacagatttcacactcaagatcagcaggggtggaggctgaggat  
ctgggagtttattactgc (SEQ ID NO:1378)

(amino acids)

GVPDRFSGSGSGTDFTLKISRVEAEDLGVYYC (SEQ ID NO:1379)

Mouse 5C6F3 light chain variable *complementarity determining regions 3 (CDR3)*  
sequence:

(DNA)

tttcaagattcacatgttcctctcacg (SEQ ID NO:1380)

(amino acids)

FQDSHVPLT (SEQ ID NO:1381)

Mouse 5C6F3 light chain variable *framework 4 (FR4)* sequence:

(DNA)

ttcgggtgctgggaccaagctggagctgaaa (SEQ ID NO:1382)

(amino acids)

FGAGTKLELK (SEQ ID NO:1383)

mu5C6F3 scFv sequence

(DNA)

gaagtgatgctggtggagtcctgggggaggcttagtgaagcctggaggggtccctgaaactctcctgtgcagcctctgga  
ttcacttttcagtacctatgccatgtcttgggttcgccagactccggagaagaggctggagtggtgcagccattag  
aatggtggtggttacacctactatccagacagtcctgaagggcgattcaccatctccagagacaatgccagaacacc  
ctgtacctgcaaatgagcagtcctgaggtctgaggacacggccacgtattaactgtgcaagacgttactatgatcactac  
tttgactactggggccaaggcaccgctctcacggctctcctcaggtggcggaggatctggcggagggtggaagcggcgga  
ggcggatccgatgttttgatgacccaaactccactctccctgacctgtcagtccttgagatcaagcctccatctcttgc  
agatctagtcagaccattgtacatagtaatggaaacacctatttagaatggtacctgcagaaaccaggccagtcctcca  
aagtcctgatctacaaagtttccaaccgattttctgggggtcccagacaggttcagtgccagtgatcagggacagat  
ttcacactcaagatcagcaggggtggaggctgaggatctgggagtttattactgctttcaagattcacatgttcctctc  
acgttcggtgctgggaccaagctggagctgaaa (SEQ ID NO:1384)

(amino acids)

EVMLVESGGGLVKPGGSLKLSAASGFTFSTYAMSWVRQTPEKRLEWVAALISNGGGYTYYPDSLKGRFTISRDN  
AKN  
TLYLQMSLSRSEDATYYCARRYDHYFDYWGGTALTIVSSGGGGSGGGGSDVLTQTPLSLPVSLGDAQSI  
SCRSSQTIVHSNGNTYLEWYLQKPGQSPKLLIYKVSNRFSVPDRFSGSGSGTDFTLKISRVEAEDLGVYYCFQDSH  
VPLTFGAGTKLELK (SEQ ID NO:1385)

3C2B1

Mouse 3C2B1 heavy chain variable *framework 1 (CDR1)* sequence:

(DNA)

gaagtgatgctggtggagtcctgggggaggcttagtgaagcctggaggggtccctgaaactctcctgtgcagcctctgga  
atcactttcagt (SEQ ID NO:1386)

(amino acids)

EVMLVESGGGLVKPGGSLKLSAASGITFS (SEQ ID NO:1387)

Mouse 3C2B1 heavy chain variable *complementarity determining regions 1 (CDR1)*  
sequence:

(DNA)

acctataccatgtcg (SEQ ID NO:1388)

(amino acids)

TYTMS (SEQ ID NO:1389)

Mouse 3C2B1 heavy chain variable *framework 2 (CDR2)* sequence:

(DNA)



tgggttcgccagactccggagaagaggctggagtggtcgca (SEQ ID NO:1390)  
(amino acids)  
WVRQTPEKRLEWVA (SEQ ID NO:1391)

Mouse 3C2B1 heavy chain variable *complementarity determining regions 2 (CDR2)* sequence:  
(DNA)  
accattagtactggtggtgataaaacctactattcagacagtggtgaaggggt (SEQ ID NO:1392)  
(amino acids)  
TISTGGDKTYYSVKG (SEQ ID NO:1393)

Mouse 3C2B1 heavy chain variable *framework 3 (CDR3)* sequence:  
(DNA)  
cgattcaccatctccagagacaatgccagaacaacctgtacctccaaatgagcagtcctgaggtctgaggacacggcc  
ttgtattactgtgcaagg (SEQ ID NO:1394)  
(amino acids)  
RFTISRDNAKNNLYLQMSSLRSEDTALYYCAR (SEQ ID NO:1395)

Mouse 3C2B1 heavy chain variable *complementarity determining regions 3 (CDR3)* sequence:  
(DNA)  
ggaaccacggctatgtattactatgctatggactac (SEQ ID NO:1396)  
(amino acids)  
GTTAMYYYAMDY (SEQ ID NO:1397)

Mouse 3C2B1 heavy chain variable *framework 4 (CDR4)* sequence:  
(DNA)  
tgggtcaaggaacctcagtcacgtctctctca (SEQ ID NO:1398)  
(amino acids)  
WGQGTSVTVSS (SEQ ID NO:1399)

Mouse 3C2B1 light chain variable *framework 1 (FW1)* sequence:  
(DNA)  
gacattgtgctgacacagtcctctgcttccttagctgtatctctggggcagagggccaccatctcatgc (SEQ ID NO:1400)  
(amino acids)  
DIVLTQSPASLAVSLGQRATISC (SEQ ID NO:1401)

Mouse 3C2B1 light chain variable *complementarity determining regions 1 (CDR1)* sequence:  
(DNA)  
agggccagcaaaagtatcagtacatctgactataattatattcac (SEQ ID NO:1402)  
(amino acids)  
RASKSISTSDYNYIH (SEQ ID NO:1403)

Mouse 3C2B1 light chain variable *framework 2 (FW2)* sequence:  
(DNA)  
tggtaccaacagaaaccaggacagccacccaaactcctcatctat (SEQ ID NO:1404)  
(amino acids)  
WYQQKPGQPPKLLIY (SEQ ID NO:1405)

Mouse 3C2B1 light chain variable *complementarity determining regions 2 (CDR2)* sequence:  
(DNA)  
CTTGCATCCAACCTAGAATCT (SEQ ID NO:1406)  
(amino acids)  
LASNLES (SEQ ID NO:1407)

Mouse 3C2B1 light chain variable *framework 3 (FW3)* sequence:

(DNA)

gggtccctgccaggttcagtggcagtggtctgggacagacttcaccctcaacatccatcctgtggaggaagaagatg  
ctgcaacctattactgt (SEQ ID NO:1408)

(amino acids)

GVPARFSGSGSGTDFTLNHPVEEEDAATYYC (SEQ ID NO:1409)

Mouse 3C2B1 light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

cagcacagtagggagcttcctctcacg (SEQ ID NO:1410)

(amino acids)

QHSRELPLT (SEQ ID NO:1411)

Mouse 3C2B1 light chain variable *framework 4 (FW4)* sequence:

(DNA)

ttcgggtgctgggaccaagctggagctgaaa (SEQ ID NO:1412)

(amino acids)

FGAGTKLELK (SEQ ID NO:1413)

Mouse B12 heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

Caggcgcagctgaaggagtcaggacctggcctgggtggcgccctcacagagcctgtccatcacttgcaactgtctctggg  
ttttcattaacc (SEQ ID NO:1414)

(amino acids)

QAQLKESGPGLVAPSQSLSTCTVSGFSLT (SEQ ID NO:1415)

Mouse B12 heavy chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

agctatggtgtacac (SEQ ID NO:1416)

(amino acids)

SYGVH (SEQ ID NO:1417)

Mouse B12 heavy chain variable *framework 2 (FW2)* sequence:

(DNA)

tgggttcgccagcctccaggaaaggtctggagtggctggga (SEQ ID NO:1418)

(amino acids)

WVRQPPGKGLEWLG (SEQ ID NO:1419)

Mouse B12 heavy chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

gtaatatggcctgggtggaagcacaaattataattcgactctcatgtccagaatg (SEQ ID NO:1420)

(amino acids)

VIWPGGSTNYNSTLMSRM (SEQ ID NO:1421)

Mouse B12 heavy chain variable *framework 3 (FW3)* sequence:

(DNA)

cggatcatcaaagacaactccaagagccaagttttcttaaaaatgaacagtcctgcaaattgatgacacagccatgtac  
tactgtgccaga (SEQ ID NO:1422)

(amino acids)

RIIKDNSKSQVFLKMNSLQIDDTAMYYCAR (SEQ ID NO:1423)

Mouse B12 heavy chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

gatcggacacctcgggtgggggcctggtttgcttac (SEQ ID NO:1424)

(amino acids)

DRTPRVGAWFAY (SEQ ID NO:1425)

Mouse B12 heavy chain variable *framework 4 (FW4)* sequence:

(DNA)

tggggccaagggactctggtcactgtctctgcag (SEQ ID NO:1426)

(amino acids)

WGQGTTLVTVSA (SEQ ID NO:1427)

Mouse B12 light chain variable *framework 1 (FR1)* sequence:

(DNA)

atcattgtgctgaccaatctccagcttctttggctgtgtctctagggcagagggccaccatatacctgc (SEQ ID NO:1428)

(amino acids)

IIVLTQSPASLAVSLGQRATISC (SEQ ID NO:1429)

Mouse B12 light chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

agagccagtgaagagtgttgctacttatggcaataattttatgcag (SEQ ID NO:1430)

(amino acids)

RASESVATYGNFMQ (SEQ ID NO:1431)

Mouse B2 heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

gaagtgggtgctggtggagtctgggggaggcttagtgagagcctggaggggtccctgaaactctcctgtgtagcctctgga  
ttcgctttcagt (SEQ ID NO:1432)

(amino acids)

EVVLVESGGGLVEPGGSLKLSVASGFAFS (SEQ ID NO:1433)

Mouse B2 heavy chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

acctttgccatgtct (SEQ ID NO:1434)

(amino acids)

TFAMS (SEQ ID NO:1435)

Mouse B2 heavy chain variable *framework 2 (FW2)* sequence:

(DNA)

tggattcgccagactccggagaagaggctggagtgggtcgca (SEQ ID NO:1436)

(amino acids)

WIRQTPEKRLEWVA (SEQ ID NO:1437)

Mouse B2 heavy chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

gccattagtaatggtggtgttacacctactatccagacactctgaagggg (SEQ ID NO:1438)

(amino acids)

AINSGGGYTYYPDTLKG (SEQ ID NO:1439)

Mouse B2 heavy chain variable *framework 3 (FW3)* sequence:

(DNA)

cgattctccatctccagagacaatgccagaataccctgtacctgcaaatgagtagtctgaggtctgaggacacggcc  
gtgtattactgtgcaaga (SEQ ID NO:1440)

(amino acids)

RFSISRDNAKNTLYLQMSSLRSEDTAVYYCAR (SEQ ID NO:1441)

Mouse B2 heavy chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

cgctactatgatctctactttgactta (SEQ ID NO:1442)

(amino acids)

RYYDLYFDL (SEQ ID NO:1443)

Mouse B2 heavy chain variable *framework 4 (FW4)* sequence:

(DNA)

Tggggccgaggcacctctctcatagtctcctca (SEQ ID NO:1444)

(amino acids)

WGRGTS LIVSS (SEQ ID NO:1445)

Mouse B2 light chain variable *framework 1 (FR1)* sequence:

(DNA)

gatattctgatgacccaaactccactctccctgctgtcagttcttgagatcaagcctccattttcttgc (SEQ ID NO:1446)

(amino acids)

DILMTQTPLSLPVS LGDQASISC (SEQ ID NO:1447)

Mouse B2 light chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

agatctagtcagaacattgtacatagtaatggaaacacctatttagaa (SEQ ID NO:1448)

(amino acids)

RSSQNIVHSNGNTYLE (SEQ ID NO:1449)

Mouse B2 light chain variable *framework 2 (FR2)* sequence:

(DNA)

tggtacctgcagaaaccaggccagtcctccaaagctcctgatctac (SEQ ID NO:1450)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:1451)

Mouse B2 light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

aaagtttccaaccgattttct (SEQ ID NO:1452)

(amino acids)

KVSNRFS (SEQ ID NO:1453)

Mouse B2 light chain variable *framework 3 (FR3)* sequence:

(DNA)

ggggtccccgcaggttcagtggttagtgggtcagggacagatttcacactcaagatcagcagagtgaggagctgaggatctgggagtttattactgc (SEQ ID NO:1454)

(amino acids)

GVPDRFSGSGSGTDFTLKISRVEAEDLGVYYC (SEQ ID NO:1455)

Mouse B2 light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

tttcaagattcacatgttcctctcacg (SEQ ID NO:1456)

(amino acids)

FQDSHVPLT (SEQ ID NO:1457)

Mouse B2 light chain variable *framework 4 (FR4)* sequence:

(DNA)

ttcgggtgctgggaccaggctggagctgaaa (SEQ ID NO:1458)  
(amino acids)  
FGAGTRLELK (SEQ ID NO:1459)

Mouse B7 heavy chain variable *framework 1 (FW1)* sequence:

(DNA)  
gaggtgcaggtggtggagctctgggggagacttagtgaagcctggagggtccctgaaactctcctgtgcagcctctgga  
ttcactttcagt (SEQ ID NO:1460)  
(amino acids)  
EVQVVESGGDLVKPGGSLKLSCAASGFTFS (SEQ ID NO:1461)

Mouse B7 heavy chain variable *complementarity determining regions 1 (CDR1)*  
sequence:

(DNA)  
agatatggcatgtct (SEQ ID NO:1462)  
(amino acids)  
RYGMS (SEQ ID NO:1463)

Mouse B7 heavy chain variable *framework 2 (FW2)* sequence:

(DNA)  
tgggttcgccagactccagacaagaggctggagtggtcgca (SEQ ID NO:1464)  
(amino acids)  
WVRQTPDKRLEWVA (SEQ ID NO:1465)

Mouse B7 heavy chain variable *complementarity determining regions 2 (CDR2)*  
sequence:

(DNA)  
accattagtagtggtggtacttacatctactatccagacagtgtgaagggg (SEQ ID NO:1466)  
(amino acids)  
TISSGGTYIYPDSVKG (SEQ ID NO:1467)

Mouse B7 heavy chain variable *framework 3 (FW3)* sequence:

(DNA)  
cgattcaccatctccagagacaatgccaaagacacctgtacctgcaaagtgcagctctgaagctctgaggacacagcc  
atgtattactgtgcaagg (SEQ ID NO:1468)  
(amino acids)  
RFTISRDNAKNTLYLQMSSLKSEDTAMYYCAR (SEQ ID NO:1469)

Mouse B7 heavy chain variable *complementarity determining regions 3 (CDR3)*  
sequence:

(DNA)  
gataactacggtagtagctacgactatgctatggactac (SEQ ID NO:1470)  
(amino acids)  
DNYGSSYDYAMDY (SEQ ID NO:1471)

Mouse B7 heavy chain variable *framework 4 (FW4)* sequence:

(DNA)  
tggggtcaaggaacctcagtcaccgtctcctca (SEQ ID NO:1472)  
(amino acids)  
WQGTSVTVSS (SEQ ID NO:1473)

Mouse B7 light chain variable *framework 1 (FR1)* sequence:

(DNA)  
gatgttttgatgacccaaactccactctccctgctgtcagttttggagatcaagcctccatctcttgc (SEQ ID  
NO:1474)  
(amino acids)  
DVLMTQTPLSLPVSLGDQASISC (SEQ ID NO:1475)

Mouse B7 light chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

agatctagtcagaccattgtacatagtaatggaaacacctatttagaa (SEQ ID NO:1476)

(amino acids)

RSSQTIVHSNGNTYLE (SEQ ID NO:1477)

Mouse B7 light chain variable *framework 2 (FR2)* sequence:

(DNA)

tggtacctgcaaaaaccaggccagtctccaaagctcctgatctac (SEQ ID NO:1478)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:1479)

Mouse B7 light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

aaagtttccaaccgattttct (SEQ ID NO:1480)

(amino acids)

KVSNRFS (SEQ ID NO:1481)

Mouse B7 light chain variable *framework 3 (FR3)* sequence:

(DNA)

gggggtcccagacaggttcagtggtggtggtcagggacagatttcacactcaagatcagcaggggtggaggctgaggatctgggagtttattactgc (SEQ ID NO:1482)

(amino acids)

GVPDRFSGSGSGTDFTLKISRVEAEDLGVYYC (SEQ ID NO:1483)

Mouse B7 light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

tttcaagattcacatgttcctctcacg (SEQ ID NO:1484)

(amino acids)

FQDSHVPLT (SEQ ID NO:1485)

Mouse B7 light chain variable *framework 4 (FR4)* sequence:

(DNA)

ttcgggtgctgggaccaagctggagctgaaa (SEQ ID NO:1486)

(amino acids)

FGAGTKLELK (SEQ ID NO:1487)

Mouse 8C7F3 heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

gaagtgatgctggtggagtcctgggggaggttagtgaagcctggaggggtccctgaaactctcctgtgcagcctctgga

ttcactttcagt (SEQ ID NO:1488)

(amino acids)

EVMLVESGGGLVKPGGSLKLSAASGFTFS (SEQ ID NO:1489)

Mouse 8C7F3 heavy chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

acctatgccatgtct (SEQ ID NO:1490)

(amino acids)

TYAMS (SEQ ID NO:1491)

Mouse 8C7F3 heavy chain variable *framework 2 (FW2)* sequence:

(DNA)

tgggttcgccagactccggagaagaggctggagtgggcgca (SEQ ID NO:1492)  
(amino acids)  
WVRQTPEKRLEWVA (SEQ ID NO:1493)

Mouse 8C7F3 heavy chain variable *complementarity determining regions 2 (CDR2)*  
sequence:  
(DNA)  
gccattagtaatggtggtggttacacctactatccagacagtctgaagggg (SEQ ID NO:1494)  
(amino acids)  
AISNGGGYTYYPDSLKG (SEQ ID NO:1495)

Mouse 8C7F3 heavy chain variable *framework 3 (FW3)* sequence:  
(DNA)  
cgattcaccatctccagagacaatgccagaacaccctgtacctgcaaatgagcagtctgaggtctgaggacacggcc  
acgtattactgtgcaaga (SEQ ID NO:1496)  
(amino acids)  
RFTISRDNAKNTLYLQMSSLRSEDTATYYCAR (SEQ ID NO:1497)

Mouse 8C7F3 heavy chain variable *complementarity determining regions 3 (CDR3)*  
sequence:  
(DNA)  
cgttactatgatcactactttgactac (SEQ ID NO:1498)  
(amino acids)  
RYYDHYFDY (SEQ ID NO:1499)

Mouse 8C7F3 heavy chain variable *framework 4 (FW4)* sequence:  
(DNA)  
tggggccaaggcaccgctctcacggtctctca (SEQ ID NO:1500)  
(amino acids)  
WQGTALTIVSS (SEQ ID NO:1501)  
Mouse 8C7F3 light chain variable *framework 1 (FR1)* sequence:  
(DNA)  
atcattgtgctgaccaatctccagcttctttggctgtgtctctagggcagagggccaccatatactgc (SEQ ID  
NO:1502)  
(amino acids)  
IIVLTQSPASLAVSLGQRATISC (SEQ ID NO:1503)

Mouse 8C7F3 light chain variable *complementarity determining regions 1 (CDR1)*  
sequence:  
(DNA)  
agagccagtgagagtgttgctacttatggcaataattttatgcag (SEQ ID NO:1504)  
(amino acids)  
RASESVATYGNNFMQ (SEQ ID NO:1505)

Mouse 8C7F3 light chain variable *framework 2 (FR2)* sequence:  
(DNA)  
tggtatcagcagaaaccaggacagccacccaaactcctcatctat (SEQ ID NO:1506)  
(amino acids)  
WYQQKPGQPPKLLIY (SEQ ID NO:1507)

Mouse 8C7F3 light chain variable *complementarity determining regions 2 (CDR2)*  
sequence:  
(DNA)  
cttgcatccaccctagattct (SEQ ID NO:1508)  
(amino acids)  
LASTLDS (SEQ ID NO:1509)

Mouse 8C7F3 light chain variable *framework 3 (FR3)* sequence:

(DNA)

ggggtccctgccaggttcagtggcagtggtctaggacagacttcaccctcaccattgatcctgtggaggctgatgat  
gctgcaacctattactgt (SEQ ID NO:1510)

(amino acids)

GVPARFSGSGSRTDFTLTIDPVEADDAATYYC (SEQ ID NO:1511)

Mouse 8C7F3 light chain variable *complementarity determining regions 3 (CDR3)*  
sequence:

(DNA)

cagcaaaataatgaggatcctccgacg (SEQ ID NO:1512)

(amino acids)

QQNNEDPPT (SEQ ID NO:1513)

Mouse 8C7F3 light chain variable *framework 4 (FR4)* sequence:

(DNA)

ttcgggtggaggcaccaagctggaaatcaag (SEQ ID NO:1514)

(amino acids)

FGGGTKLEIK (SEQ ID NO:1515)



Mouse H11 heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

gaagtgggtgctgggtggagctctgggggaggccttagtggagcctggagggtccctgaaactctcctgtgtagcctctgga  
ttcgcttttagt (SEQ ID NO:1516)

(amino acids)

EVVLVESGGGLVEPGGSLKLSCVASGFAFS (SEQ ID NO:1517)

Mouse H11 heavy chain variable *complementarity determining regions 1 (CDR1)*  
sequence:

(DNA)

acctttgccatgtct (SEQ ID NO:1518)

(amino acids)

TFAMS (SEQ ID NO:1519)

Mouse H11 heavy chain variable *framework 2 (FW2)* sequence:

(DNA)

tggattcgccagactccggagaagaggtggagtgggtcgca (SEQ ID NO:1520)

(amino acids)

WIRQTPEKRLEWVA (SEQ ID NO:1521)

Mouse H11 heavy chain variable *complementarity determining regions 2 (CDR2)*  
sequence:

(DNA)

gccattagtaatggtggtggttacacttactatccagacactctgaagggg (SEQ ID NO:1522)

(amino acids)

AISNGGGYTYYPDTLKG (SEQ ID NO:1523)

Mouse H11 heavy chain variable *framework 3 (FW3)* sequence:

(DNA)

cgattcaccatctccagagacaatgccagaataccctgtacctgcaaatgagtagtctgaggtctgaggacacggcc  
gtgtattactgtgcaaga (SEQ ID NO:1524)

(amino acids)

RFTISRDNAKNTLYLQMSSLRSEDTAVYYCAR (SEQ ID NO:1525)

Mouse H11 heavy chain variable *complementarity determining regions 3 (CDR3)*  
sequence:

(DNA)

cgctactatgatctctactttgactta (SEQ ID NO:1526)

(amino acids)

RYYDLYFDL (SEQ ID NO:1527)

Mouse H11 heavy chain variable *framework 4 (FW4)* sequence:

(DNA)

tggggccaaggcacctctctcatagtctcctca (SEQ ID NO:1528)

(amino acids)

WGQGTSLIVSS (SEQ ID NO:1529)

Mouse H11 light chain variable *framework 1 (FR1)* sequence:

(DNA)

gatattctgatgacccaaactccactctccctgcctgtcagtccttggagatcaagcctccattttcttgc (SEQ ID  
NO:1530)

(amino acids)

DILMTQTPLSLPVSLGDQASISC (SEQ ID NO:1531)

Mouse H11 light chain variable *complementarity determining regions 1 (CDR1)*  
sequence:

(DNA)

agatctagtcagaacattgtacatagtaatggaaacacctatttagaa (SEQ ID NO:1532)

(amino acids)

RSSQNIVHSNGNTYLE (SEQ ID NO:1533)

Mouse H11 light chain variable *framework 2 (FR2)* sequence:

(DNA)

tggtacctgcagaaaccaggccagtcctccaaagctcctgatctac (SEQ ID NO:1534)

(amino acids)

WYLQKPGQSPKLLIY (SEQ ID NO:1535)

Mouse H11 light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

aaagtttccaaccgattttct (SEQ ID NO:1536)

(amino acids)

KVSNRFS (SEQ ID NO:1537)

Mouse H11 light chain variable *framework 3 (FR3)* sequence:

(DNA)

gggggtccccgcagaggttcagtggttagtgggtcagggacagatttcacactcaagatcagcagagtgaggagctgaggat  
ctgggagtttattactgc (SEQ ID NO:1538)

(amino acids)

GVPDRFSGSGSGTDFTLKISRVEAEDLGVYYC (SEQ ID NO:1539)

Mouse H11 light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

tttcaagattcacatgttcctctcacg (SEQ ID NO:1540)

(amino acids)

FQDSHVPLT (SEQ ID NO:1541)

Mouse H11 light chain variable *framework 4 (FR4)* sequence:

(DNA)

ttcgggtgctgggaccaggctggagctgaaa (SEQ ID NO:1542)

(amino acids)

FGAGTRLELK (SEQ ID NO:1543)

Mouse B9 heavy chain variable *framework 1 (FW1)* sequence:

(DNA)

gaggtgcaggtggtggagtcctgggggagacttagtgaagcctggagggtccctgaaactctcctgtgcagcctctgga  
ttcactttcagt (SEQ ID NO:1544)

(amino acids)

EVQVVESGGDLVKPGGSLKLSAASGFTFS (SEQ ID NO:1545)

Mouse B9 heavy chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

agatatggcatgtct (SEQ ID NO:1546)

(amino acids)

RYGMS (SEQ ID NO:1547)

Mouse B9 heavy chain variable *framework 2 (FW2)* sequence:

(DNA)

tgggttcgccagactccagacaagaggctggagtggtcgca (SEQ ID NO:1548)

(amino acids)

WVRQTPDKRLEWVA (SEQ ID NO:1549)

Mouse B9 heavy chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

accattagtagtggtggtacttacatctactatccagacagtggtgaagggg (SEQ ID NO:1550)

(amino acids)

TISSGGTYIYPDSVKG (SEQ ID NO:1551)

Mouse B9 heavy chain variable *framework 3 (FW3)* sequence:

(DNA)

cgattcaccatctccagagacaatgccagaacacccctgtacctgcaaattgagcagttctgaagttctgaggacacagcc

atgtattactgtgca (SEQ ID NO:1552)

(amino acids)

RFTISRDNAKNTLYLQMSSLKSEDTAMYICAR (SEQ ID NO:1553)

Mouse B9 heavy chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

agggataactacggtagtagctacgactatgctatggactac (SEQ ID NO:1554)

(amino acids)

DNYGSSYDYAMDY (SEQ ID NO:1555)

Mouse B9 heavy chain variable *framework 4 (FW4)* sequence:

(DNA)

tggggtcaaggaacctcagtcaccgtctcctct (SEQ ID NO:1556)

(amino acids)

WGQGTSTVTVSS (SEQ ID NO:1557)

Mouse B9 light chain variable *framework 1 (FR1)* sequence:

(DNA)

caaattgtttctcaccagttctccagcaatcatgtctgcattctccaggggaggagggtcaccctaacctgc (SEQ ID NO:1558)

(amino acids)

QIVLTQSPAIMASAPGEEVTLTC (SEQ ID NO:1559)

Mouse B9 light chain variable *complementarity determining regions 1 (CDR1)* sequence:

(DNA)

agtgccagctcaagtgttaagttacatgcac (SEQ ID NO:1560)

(amino acids)

SASSSVSYMH (SEQ ID NO:1561)

Mouse B9 light chain variable *framework 2 (FR2)* sequence:

(DNA)

tggttcagcagaggccaggcacttctccaaactctggatttat (SEQ ID NO:1562)

(amino acids)

WFQQRPGTSPKLWIY (SEQ ID NO:1563)

Mouse B9 light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

accacatccaacctggcttct (SEQ ID NO:1564)

(amino acids)

TTSNLAS (SEQ ID NO:1565)

Mouse B9 light chain variable *framework 3 (FR3)* sequence:

(DNA)

ggagtcacctgctcgcttcagtgaggcagtgatctgggacctcttactctctcacaatcagccgaatggaggctgaagat  
gctgccacttattactgc (SEQ ID NO:1566)

(amino acids)

GVPARFSGSGSGTSYSLTISRMEAEDAATYYC (SEQ ID NO:1567)

Mouse B9 light chain variable *complementarity determining regions 3 (CDR3)*  
sequence:

(DNA)

cagcaaaggagtagttacccattc (SEQ ID NO:1568)

(amino acids)

QQRSSYPF (SEQ ID NO:1569)

Mouse B9 light chain variable *framework 4 (FR4)* sequence:

(DNA)

acgttcggctcggggacaaagttggaaataaaa (SEQ ID NO:1570)

(amino acids)

TFSGGTKLEIK (SEQ ID NO:1571)

mu3C2B1 scFv sequence

(DNA)

Gaagtgatgctggtggagtcctgggggaggcttagtgaagcctggagggtccctgaaactctcctgtgcagcctctgga  
atcactttcagtagctataccatgtcgtgggttcgccagactccggagaagaggctggagtgggtcgcaaccattagt  
actggtggtgataaaacctactattcagacagtggtgaagggtcgattcaccatctccagagacaatgccagaacaac  
ctgtacctccaaatgagcagtcctgaggtctgaggacacggccttgattactgtgcaaggggaaccacggctatgtat  
tactatgctatggactactgggtcaaggaacctcagtcacgcgtctcctcaggtggcggaggatctggcggagggtgga  
agcggcggaggcggatccgacattgtgctgacacagtcctcctgcttccttagctgtatctctggggcagagggccacc  
atctcatgcagggccagcaaaagtatcagtagctctgactataattatattcactggtaccaacagaaaccaggacag  
ccacccaaactcctcatctatcttgcatccaacctagaatctgggggtccctgccaggttcagtggcagtggtctggg  
acagacttcacctcaacatccatcctgtggaggagaagatgctgcaacctattactgtcagcacagtagggagctt  
cctctcaggttcgggtgctgggaccaagctggagctgaaa (SEQ ID NO:1572)

(amino acids)

EVMLVESGGGLVPGGSLKLSAASGITFSTYTMWVRQTPEKRLEWVATISTGGDKTYSDSVKGRFTISRDNNAKN  
NLYLQMSLRSEDTALYYCARGTTAMYYYAMDYWGQTSVTVSSGGGGSGGGGSDIVLTQSPASLAVSLGQR  
ATISCRASKSISTSDYNIHWYQQKPGQPPLLIYLASNLESGVPARFSGSGSGTDFTLNIHPVEEEDAATYYCQHS  
RELPLTFGAGTKLELK (SEQ ID NO:1573)

mu20A10 scFv full sequence

(DNA)

gaagtgatgctggtggaatctggcggcggactgggttaagcctggcggatctctgaagctgagctgtgccgccagcggc  
ttcaccttttagcacatacgccatgagctggatccggcagacccctgagaagagactggaatgggttgccagcatcggc  
agagccggcagcacctactacagcgattctgtgaagggcagattcaccatcagccgggacaacgtgcggaacatcctg  
tacctgcagatgagcagcctgcggagcggagataccgccatgtactactgtgccagaggacccatctacaacgactac  
gacgagttcgccatttggggccagggcacactgggttacagtttctgctggtggcggaggatctggcggagggtggaagc  
ggcggaggcggatccaatatcatgatgacacagagccccagcagcctggctgtgtctgctggcgagaaagtgacctg  
tctgcaagagcagccagagcgtgctgtactccagcaaccagaagaactacctggcctggtatcagcagaagcccggc  
cagtcctctaagctgctgactactgggccagcaccagagaaagcggcgtgcccgatagattcacaggcagcggcagc  
ggaaccgacttcacctgacaatcagctctgtgcaggccgaagatctggcgtgtactattgccaccagtagcctgtcc  
agcctgacctttggcgccggaacaaagctggaactgaag (SEQ ID NO:1574)

(amino acids)

EVMLVESGGGLVPGGSLKLSAASGFTFSTYAMSWIRQTPEKRLEWVASIGRAGSTYYSDSVKGRFTISRDNVRNI  
LYLQMSLRSEDTAMYYCARGPIYNDYDEFAYWGQGLTVTSAGGGGSGGGGSGGGGSDIVLTQSPSSSLAVSAGEKV  
TMSCKSSQSVLYSSNQKNYLAWYQQKPGQSPKLLIYWASTRESGVPDRFTGSGSGTDFTLTISVQAEDLAVYYCHQ  
YLSSLTFGAGTKLELK (SEQ ID NO:1575)

hu20A10M scFV

(DNA)

gaggtgcagctggttgaatctggcgggcggaacttgtgaagcctggcggaatctctgagactgagctgtgccgccagcggc  
ttcaccttttagcacatacgccatgagctgggtccgacaggccccctggaaaaggccttgaatgggttgccctctatcggc  
agagccggcagcacctactacagcgattctgtgaagggcagattcaccatcagccgggacaacgccaagaacagcctg  
tacctgcagatgaactccctgagagccgaggacaccgcccgtgtactattgtgccagaggacccatctacaacgactac  
gacgagttcgccctattggggccagggcacactgggtcacagtcagctctggcggtggcggaagcggaggcggtggctcc  
ggtggcgaggagcagcgacatcgtgatgacacagagcccttctagcctggcgcgtgtctctgggagagagagccacaatc  
agctgcaagagcagccagagcgtgctgtactccagcaaccagaagaactacctggcctgggtatcagcagaagccccgga  
cagtcctcccaagctgctgatctactgggcccagcaccagagaaaaggcggtgcccgatagattcacaggctctggcagc  
ggcaccgacttcacctgacaattagcagtcctgcaggccgaggacgtggcgcgtgtactactgtcaccagtacctgagc  
agcctgacctttggcgggcggaacaaaggtggaaatcaag (SEQ ID NO:1576)

(amino acids)

EVQLVESGGGLVKGPGSLRLSCAASGFTFTSTYAMSWVRQAPGKGLEWVASIGRAGSTYYSDSVKGRFTISRDNKNS  
LYLQMNSLRAEDTAVYYCARGPIYNDYDEFAYWGQGLTVTVSSGGGGSGGGSGGGSDIVMTQSPSSLAVSLGERA  
TISCKSSQSVLYSSNQKNYLAWYQQKPGQSPKLLIYWASTRESGVPDRFTGSGSGTDFTLTISLQAEDVAVYYCHQ  
YLSSLTFGGGTKVEIK (SEQ ID NO:1577)

hu20A10C2 scFV

(DNA)

gaggtgcagctggttgaatctggcgggcggaacttgtgaagcctggcggaatctctgagactgagctgtgccgccagcggc  
ttcaccttttagcacatacgccatgagctgggtccgacaggccccctggaaaaggccttgaatgggttgccctctatcggc  
agagccggcagcacctactacagcgattctgtgaagggcagattcaccatcagccgggacaacgccaagaacagcctg  
tacctgcagatgaactccctgagagccgaggacaccgcccgtgtactattgtgccagaggacccatctacaacgactac  
gacgagttcgccctattggggccagggcacactgggtcacagtcagctctggcggtggcggaagcggaggcggtggctcc  
ggtggcgaggagcagcgacattgtgctgacccagtcctccagcctccttggcgcgtgtctccaggacagagggccaccatc  
acctgcaagagcagccagagcgtgctgtactccagcaaccagaagaactacctggcctgggtatcagcagaaaccagga  
caacctcctaaactcctgatttactgggcccagcaccagagaaaaggcggtgcccagcaggttcagcggcagtggtgtct  
gggaccgatttcacctcacaattaatcctgtggaagctaataactgataactgcaaattattactgtcaccagtacctgagc  
agcctgaccttcggcgaggaggaccaaggtggagatcaaacga (SEQ ID NO:1578)

(amino acids)

EVQLVESGGGLVKGPGSLRLSCAASGFTFTSTYAMSWVRQAPGKGLEWVASIGRAGSTYYSDSVKGRFTISRDNKNS  
LYLQMNSLRAEDTAVYYCARGPIYNDYDEFAYWGQGLTVTVSSGGGGSGGGSGGGSDIVLTQSPASLAVSPQRA  
TITCKSSQSVLYSSNQKNYLAWYQQKPGQPPKLLIYWASTRESGVPARFSGSGSGTDFTLTINPVEANDTANYCHQ  
YLSSLTFGGGTKVEIKR (SEQ ID NO:1579)

hu20A10N scFV

(DNA)

caggtgcagctggttgaatctggcgggcggaacttgtgaagcctggcggaatctctgagactgagctgtgccgccagcggc  
ttcaccttttagcacatacgccatgagctggatcagacaggccccctggcaaaggcctggaatgggtggcgctctattggc  
agagccggcagcacctactacagcgactctgtgaagggcagattcaccatcagccgggacaacgccaagaacagcctg  
tacctgcagatgaactccctgagagccgaggacaccgcccgtgtactattgtgccagaggacccatctacaacgactac  
gacgagttcgccctattggggccagggcacactgggtcacagtttctagcggcggtggcggaagcggaggcggtggctcc  
ggtggcgaggagcagcgaaaattgtgctgacacagagccccgccacactgtcactttctccaggcgaaagagccacactg  
agctgcaagagcagccagagcgtgctgtactccagcaaccagaagaactacctggcctgggtatcagcagaagccccggc  
caagctcctcggtgctgatctattgggcccagcacaagagagagcggcgcctcctgccagattttctggcagcggtct  
ggcaccgatttcacctgaccataagcagcctggaacctgaggacttcgcgcgtgtattactgccaccagtacctgagc  
agcctgacctttggcgagggcaccaaggtggaaatcaagcgg (SEQ ID NO:1580)

(amino acids)

QVQLVESGGGLVKGPGSLRLSCAASGFTFTSTYAMSWIRQAPGKGLEWVASIGRAGSTYYSDSVKGRFTISRDNKNS  
LYLQMNSLRAEDTAVYYCARGPIYNDYDEFAYWGQGLTVTVSSGGGGSGGGSGGGSEIVLTQSPATLSLSPGERA  
TLSCCKSSQSVLYSSNQKNYLAWYQQKPGQAPRLLIYWASTRESGIPARFSGSGSGTDFTLTISLLEPEDFAVYYCHQ  
YLSSLTFGGGTKVEIKR (SEQ ID NO:1581)

mu20A10-CAR T-8-4-1BB-3z

(DNA)

atggccttaccagtgaaccgccttgctcctgccgctggccttgetgctccaagccgccaggccggaagtgatgctgggtg  
 gaatctggcggcggaactggttaagcctggcggtatctctgaagctgagctgtgccgccagcggttcacctttagcaca  
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 (SEQ ID NO:1582)

(amino acids)

MALPVTALLLPLALLLHAARPEVMLVESGGGLVKPGGSLKLSCAASGFTFSTYAMSWIRQTPEKRLEWVASIGRAGS  
 TYYSDSVKGRFTISRDNVRNILYLQMSLRSEDAMYYCARGPIYNDYDEFAYWGQGLVTVSAGGGGSGGGGSGGG  
 GSNIMMTQSPSSLAVSAGEKVTMSCKSSQSVLYSSNQKNYLAWYQQKPGQSPKLLIYWASTRESGVPDRFTGSGSGT  
 DFTLTISVQAEDLAVYYCHQYLSSLTFGAGTKLELKTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGL  
 DFACDIYIWAPLAGTCGVLLLSLVITLYCKRGRKKLLYIFKQPFMRPVQTTQEEDGCSCRFPEEEEEGGCELRVKFSR  
 SADAPAYKQGQNLQYLNELNLGRREEYDVLDRRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRG  
 KGHDLGYQLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:1583)

hu20A10-CAR T-8-4-1BB-3z

(DNA)

atggccttaccagtgaaccgccttgctcctgccgctggccttgetgctccaagccgccaggccggagggtgcagctgggtt  
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 (SEQ ID NO:1584)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKPGGSLRLSCAASGFTFSTYAMSWVRQAPGKGLEWVASIGRAGS  
 TYYSDSVKGRFTISRDNKNSLYLQMSLRSEDAMYYCARGPIYNDYDEFAYWGQGLVTVSSGGGGSGGGGSGGG  
 GSDIVLTQSPASLAVSPGQRATITCKSSQSVLYSSNQKNYLAWYQQKPGQPPLLIYWASTRESGVPARFSGSGSGT  
 DFTLTINPVEANDTANYCHQYLSSLTFGGGTVKVEIKRTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRG  
 LDFACDIYIWAPLAGTCGVLLLSLVITLYCKRGRKKLLYIFKQPFMRPVQTTQEEDGCSCRFPEEEEEGGCELRVKFS

RSADAPAYKQGQNQLYNELNLGRREEYDVLDKRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRR  
KGHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:1585)

mu20A10-CAR T-8-28-3z

(DNA)

atggccttaccagtgaccgccttgcctcctgccgctggccttgcctgcctccacgcgcgcagggccggaagtgatgctggtg  
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cagggctctcagtacagccaccaaggacacctacgacgccttcacatgcaggccctgccccctcgctgataa (SEQ  
ID NO:1586)

(amino acids)

MALPVTALLLPLALLLHAARPEVMLVESGGGLVKPGGSLKLSCAASGFTFSTYAMSWIRQTPEKRLEWVASIGRAGS  
TYYSDSVKGRFTISRDNVRNILYLQMSLSRSEDAMYYCARGPIYNDYDEFAYWGQGLVTVSAGGGGSGGGGSGGG  
GSNIMMTQSPSSLAVSAGEKVTMSCKSSQSVLYSSNQKNYLAWYQQKPKQLLIYWASTRESGVPDRFTGSGSGT  
DFTLTISVQAEDLAVYYCHQYLSSLTFGAGTKLELKTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGL  
DFACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSRS  
ADAPAYKQGQNQLYNELNLGRREEYDVLDKRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRCK  
GHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:1587)

hu20A10-CAR T-8-28-3z

(DNA)

atggccttaccagtgaccgccttgcctcctgccgctggccttgcctgcctccacgcgcgcagggccggaagtgacgtggtt  
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(SEQ ID NO:1588)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKPGGSLRLSCAASGFTFSTYAMSWVRQAPGKGLEWVASIGRAGS  
 TYYSDSVKGRFTISRDNAKNSLYLQMNSLRAEDTAVYYCARGPIYNDYDEFAYWGQGLVTVSSGGGGSGGGSGGG  
 GSDIVLTQSPASLAVSPGQRATITCKSSQSVLYSSNQKNYLAWYQQKPGQPPKLLIYWASTRESGVPARFSGSGSGT  
 DFTLTINPVEANDTANYYCHQYLSSLTFGGGTKEIKRTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRG  
 LDFACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSR  
 SADAPAYKQGQNQLYNELNLGRREEYDVLDRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRG  
 KGHGGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:1589)

mu20A10-CAR T-8-4-1BB-3z-1XX

(DNA)

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 ttccagggtctcagtacagccaccaaggacaccttcgacgccttcacatgcagggccttccccctcgctgataa  
 (SEQ ID NO:1590)

(DNA)

MALPVTALLLPLALLLHAARPEVMLVESGGGLVKPGGSLKLSCAASGFTFSTYAMSWIRQTPEKRLEWVASIGRAGS  
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 GSNIMMTQSPSSLAVSAGEKVTMSCKSSQSVLYSSNQKNYLAWYQQKPGQSPKLLIYWASTRESGVPRFTGSGSGT  
 DFTLTISVQAEDLAVYYCHQYLSSLTFGAGTKLELKTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGL  
 DFACDIYIWAPLAGTCGVLLLSLVITLYCKRGRKKLLYIFKQPFMRPVQTTQEEDGCSRFEEEEGGCELRVKFSR  
 SADAPAYKQGQNQLYNELNLGRREEYDVLDRRGRDPEMGGKPRRKNPQELFNEQLQKDKMAEAFSEIGMKGERRRG  
 KGHGGLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1591)

hu20A10-CAR T-8-4-1BB-3z-1XX

(DNA)

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(SEQ ID NO:1592)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKPGGSLRLSCAASGFTFTSTYAMSWVRQAPGKGLEWVASIGRAGS  
TYYSDSVKGRFTISRDNKNSLYLQMNSLRAEDTAVYYCARGPIYNDYDEFAYWGQGLVTVSSGGGSGGGSGGG  
GSDIVLTQSPASLAVSPGQRATITCKSSQSVLYSSNQKNYLAWYQQKPGQPPKLLIYWASTRESGVPARFSGSGSGT  
DFTLTINPVEANDTANYYCHQYLSSLTFGGGTKVEIKRTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRG  
LDFACDIYIWAPLAGTCGVLLLSLVITLYCKRGRKKLLYIFKQPFMRPVQTTQEEDGCSCRFEEEEEGGCELRVKFS  
RSADAPAYKQGQNQLYNELNLGRREEYDVLDRRGRDPEMGGKPRRKNPQEGLFNQLQDKMAEAFSEIGMKGERRR  
KGHDGLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1593)

mu20A10-CAR T-8-28-3z-1XX

(DNA)

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ID NO:1594)

(amino acids)

MALPVTALLLPLALLLHAARPEVMLVESGGGLVKPGGSLKLSCAASGFTFTSTYAMSWIRQTPEKRLEWVASIGRAGS  
TYYSDSVKGRFTISRDNVRNIIYLQMSSLRSEDTAMYYCARGPIYNDYDEFAYWGQGLVTVSAGGGGSGGGSGGG  
GSNIMMTQSPSSLAVSAGEKVTMSCKSSQSVLYSSNQKNYLAWYQQKPGQSPKLLIYWASTRESGVPDRFTGSGSGT  
DFTLTISVQAEDLAVYYCHQYLSSLTFGAGTKLELKTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGL  
DFACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSRS  
ADAPAYKQGQNQLYNELNLGRREEYDVLDRRGRDPEMGGKPRRKNPQEGLFNQLQDKMAEAFSEIGMKGERRRGK  
GHDGLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1595)

hu20A10-CAR T-8-28-3z-1XX

(DNA)

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 (SEQ ID NO:1596)

(amino acids)

MALPVTALLLPLALLHAARPEVQLVESGGGLVKPGGSLRLSCAASGFTFSTYAMSWVRQAPGKGLEWVASIGRAGS  
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 DFTLTINPVEANDTANYCHQYLSSLTFGGGTKVEIKRTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRG  
 LDFACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSR  
 SADAPAYKQQNQLYNELNLGRREEYDVLDRGRDPEMGGKPRRKNPQEGFLFNLQKDKMAEAFSEIGMKGERRRG  
 KGHDGLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1597)

25E6

mu25E6 scFv full sequence

(DNA)

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(amino acids)

EVQLVESGGDLVKPGGSLKLSAASGFTSSYGMSWVRQTPDKRLEWVATISNGGRHTFYPDSVKGRFTISRDNKN  
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 ISCKSSQSLDSDGKTYLNWLLQRPQSPKRLIYLVSKLDSGVPDRFTGSGSGTDFTLKISRVEAEDLGYYCQWGT  
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hu25E6 full sequence

(DNA)

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(amino acids)

EVQLVESGGGLVKPGGSLRLSCAASGFTSSYGMSWVRQAPGKGLEWVSTISNGGRHTFYPDSVKGRFTISRDNKN  
 SLYLQMNSLRAEDTAVYYCARQTGTGEGWFAYWGQTLVTVSSGGGGSGGGSGGGSDIVMTQTPLSLSVTPGQPAS  
 ISCKSSQSLDSDGKTYLNWYLQKPGQSPQLLIYLVSKLDSGVPDRFSGSGSGTDFTLKISRVEAEDVGVYYCQWGT  
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mu25E6-CAR T-8-4-1BB-3z

(DNA)

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 ID NO:1602)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGDLVKPGGSLKLSAASGFTFSSYGMSWVRQTPDKRLEWVATISNGGR  
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 SDVVMQTPLTSLVTIGQPASISCKSSQSLDSDGKTYLNWLLQRPQGSPKRLIYLVSKLDSGVPDRFTGSGSGTDF  
 TLKISRVEAEDLGVIYCWQGTHTFPQTGGGKLEIKTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLD  
 FACDIYIWAPLAGTCGVLLSLVITLYCKRGRKLLYIFKQPFMRPVQTTQEEDGCSCRFEEEEGGCELRVKFSRS  
 ADAPAYKQGNQLYNELNLGRREEYDVLDRKRRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRGK  
 GHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:1603)

hu25E6-CAR T-8-4-1BB-3z

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 ID NO:1604)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKPGGSLRLSAASGFTFSSYGMSWVRQAPGKGLEWVSTISNGGR  
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 TLKISRVEAEDVGVYCWQGTHTFPQTGGGKLEIKTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLD  
 FACDIYIWAPLAGTCGVLLSLVITLYCKRGRKLLYIFKQPFMRPVQTTQEEDGCSCRFEEEEGGCELRVKFSRS

ADAPAYKQGQNQLYNELNLGRREEYDVLDRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRGK  
GHDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:1605)

mu25E6-CAR T-8-28-3z

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NO:1606)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGDLVKPGGSLKLSCAASGFTFSSYGMSWVRQTPDKRLEWVATISNGGR  
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SDVVMQTPLTSLVTIGQPASISCKSSQSLDSDGKTYLNLWLLQRPQSPKRLIYLVSKLDSGVPDRFTGSGSGTDF  
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FACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSRSA  
DAPAYKQGQNQLYNELNLGRREEYDVLDRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRGK  
HGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:1607)

hu25E6-CAR T-8-28-3z

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NO:1608)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVLPKPGSLRLSCAASGFTFSSYGMSWVRQAPGKGLEWVSTISNGGR  
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FACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSRSA  
DAPAYKQGQNQLYNELNLGRREEYDVLDRKRRGRDPEMGGKPRRKNPQEGLYNELQKDKMAEAYSEIGMKGERRRRGK  
HDGLYQGLSTATKDTYDALHMQALPPR\*\* (SEQ ID NO:1609)

mu25E6-CAR T-8-4-1BB-3z-1XX

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ID NO:1610)

(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGDLVLPKPGSLKLSAASGFTFSSYGMSWVRQTPDKRLEWVATISNGGR  
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SDVVMQTPLTLSTIGQPASISCKSSQSLDSDGKTYLNWLLQRPQGSPKRLIYLVSKLDSGVPDRFTGSGSGTDF  
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GHDGLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1611)

hu25E6-CAR T-8-4-1BB-3z-1XX

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(amino acids)

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mu25E6-CAR T-8-28-3z-1XX

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(amino acids)

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hu25E6-CAR T-8-28-3z-1XX

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NO:1616)

(amino acids)

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TLKISRVEAEDVGYYCWQGHFPTFGQGTKEIKTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLD  
FACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSRSA  
DAPAYKQGQNQLYNELNLGRREEYDVLDKRRGRDPEMGGKPRRKNPQEGFLNELQKDKMAEAFSEIGMKGERRRGKG  
HDGLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1617)

MNC2-1XX

muMNC2-CAR T-8-4-1BB-3z-1XX

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(amino acids)

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GSGGGSGGGGSDIVITQSTASLGVSLGQRATISCRASKSVSTSGYSYMHYQQRPGQPPKLLIYLASNLESGVPAR  
FSGSGSGTDFTLNIHPVEEEDAATYYCQHSRELPTFGGGTKLEIKRADAAPTSTTTPAPRPPTPAPTIASQPLSL  
RPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLLSLVITLYCKRGRKLLYIFKQPFMRPVQTTQEEDGCSC  
RFPEEEEGGCELVRVKFSRSADAPAYKQGQNQLYNELNLGRREEYDVLDKRRGRDPEMGGKPRRKNPQEGFLNELQKD  
KMAEAFSEIGMKGERRRGKGHDGLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1619)

huMNC2-CAR T-8-4-1BB-3z-1XX

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taa (SEQ ID NO:1620)

(amino acids)

MALPVTALLLPLALLHAARPEVQLVESGGGLVKGPGSLRLSCAASGFTFSGYAMSWVRQAPKGLEWVSTISSGGT  
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GGSDIVLTQSPASLAIVSPGQRATITCRASKSVSTSGYSYMHWYQQKPGQPPKLLIYLASNLESGVPAARFSGSGGTD  
FTLTINPVEANDTANYYCQHSRELPFTFGGGTKVEIKRTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTR  
GLDFACDIYIWAPLAGTCGVLLLSLVITLYCKRGRKLLYIFKQPFMRPVQTTQEEDGCSRFPEEEEGGCELRVKF  
SRSADAPAYKQGQNQLYNELNLGRREEYDVLDRGRDPEMGGKPRRKNPQEGLFNELQKDKMAEAFSEIGMKGERR  
RGKGDGLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1621)

muMNC2-CAR T-8-28-3z-1XX

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(amino acids)

MALPVTALLLPLALLHAARPEVQLEESGGGLVKGPGSLKLSCAASGFTFSGYAMSWVRQTPEKRLEWVATISSGGT  
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SGGGGSGGGGSDIVITQSTASLGVS LGQRATISCRASKSVSTSGYSYMHWYQQRPGQPPKLLIYLASNLESGVPAAR  
FSGSGSGTDFTLNIHPVEEEDAATYYCQHSRELPFTFGGGTKLEIKRADAAPT VSTTTPAPRPPTPAPTIASQPLSL  
RPEACRPAAGGAVHTRGLDFACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPY  
APPRDFAAYRSRVKF SRSADAPAYKQGQNQLYNELNLGRREEYDVLDRGRDPEMGGKPRRKNPQEGLFNELQKDK  
MAEAFSEIGMKGERRRGKGDGLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1623)

huMNC2-CAR T-8-28-3z-1XX

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 (SEQ ID NO:1624)

(amino acids)

MALPVTALLLPLALLHAARPEVQLVESGGGLVKGPGSLRLSCAASGFTFSGYAMSWVRQAPGKGLEWVSTISSGGT  
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 GGSDIVLTQSPASLAVSPGQRATITCRASKSVSTSGYSYMHWYQQKPGQPPKLLIYLASNLESGVPAARFSGSGSTG  
 FTLTINPVEANDTANYYCQHSRELPTFTGGGKVEIKRTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTR  
 GLDFACDIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFS  
 RSADAPAYKQGQNQLYNELNLGRREEYDVLDRKRRDPEMGGKPRRKNPQEGFLFNLQKDKMAEAFSEIGMKGERRR  
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MNE6-1XX

muMNE6-CAR T-8-4-1BB-3z-1XX

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 NO:1626)

(amino acids)

MALPVTALLLPLALLHAARPEVKVVESSGDLVKPGSLKLSCVVSIGFTFSRYGMSWVRQTPGKRLEWVATISSGGT  
 YIYYPDSVKGRFTISRDNKNTLYLQMSSLSKSEDTAMYHCTRDNYGRNYDYGMDYWGQGTSTVTVSSGGGSGGGGSG  
 GGSQIVLTQSPAIMSASPGEVTLTCSATSSVSIIHWFQRRPGTSPKLWIYSTNLASGVVPRFSGSGYGTSLT  
 ISRMEAEDAATYYCQRSSSPFTFGSGTKLEIKRTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFAC  
 DIYIWAPLAGTCGVLLLSLVITLYCKRGRKKLLYIFKQPFMRPVQTTQEEDGCSCRFEEEEEGGCELRVKFSRSADA  
 PAYKQGQNQLYNELNLGRREEYDVLDRKRRDPEMGGKPRRKNPQEGFLFNLQKDKMAEAFSEIGMKGERRRKGKHD  
 GLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1627)

huMNE6-CAR T-8-4-1BB-3z-1XX

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 NO:1628)

(amino acids)

MALPVTALLLPLALLHAARPEVQLVESGGGLVKGPGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGT  
 YIYYPDSVKGRFTISRDNKNTLYLQMNSLRAEDTAVYYCTRDNYGRNYDYGMDYWGQGLVTVSSGGGGSGGGGSG  
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 PAYKQGQNQLYNELNLGRREEYDVLDRRRGRDPEMGGKPRRKNPQEGFLNELQKDKMAEAFSEIGMKGERRRGKGHD  
 GLFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1629)

muMNE6-CAR T-8-4-28-3z-1XX

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(amino acids)

MALPVTALLLPLALLHAARPEVKVVESSGDLVKPGGSLKLSCVVSIGFTFSRYGMSWVRQTPGKRLEWVATISGGGT  
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 GGSQIVLTQSPAIMSASPGEVTLTCSATSSVSYIHWYQQRPGTSPKLWIYSTSNLASGVPVRFSGSGYGTYSYSLT  
 ISRMEAEDAATYYCQQRSSSPFTFGSGTKLEIKTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFAC  
 DIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSRSADAP  
 AYKQGQNQLYNELNLGRREEYDVLDRRRGRDPEMGGKPRRKNPQEGFLNELQKDKMAEAFSEIGMKGERRRGKGHDG  
 LFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1631)

huMNE6-CAR T-8-4-28-3z-1XX

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(amino acids)

MALPVTALLLPLALLLHAARPEVQLVESGGGLVKGPGSLRLSCAASGFTFSRYGMSWVRQAPGKRLEWVSTISGGGT  
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GGGSEIVLTQSPATLSLSPGERATLTCSATSSVSYIHQYQQRPGQSPRLLIYSTSNLASGIPARFSGSGSGSDYTLT  
ISSLEPEDFAVYYCQQRSSSPFTFGSGTKVEIKTTTPAPRPPTPAPTIASQPLSLRPEACRPAAGGAVHTRGLDFAC  
DIYIWAPLAGTCGVLLLSLVITLYCRSKRSRLHSDYMNMTPRRPGPTRKHYPYAPPRDFAAYRSRVKFSRSADAP  
AYKQGQNQLYNELNLGRREEYDVLDKRRGRDPEMGGKPRRKNPQEGFLNELQKDKMAEAFSEIGMKGERRRGKGHDC  
LFQGLSTATKDTFDALHMQALPPR\*\* (SEQ ID NO:1633)

Minimal CMV promoter (mCMV)  
(DNA)

taggcgtgtacggtgggaggcctatataagcagagctcggttagtgaaccgtcagatcgctggagacgccatccacg  
ctgttttgacctccatagaagacaccgggaccgatccag (SEQ ID NO:1634)

Minimal IL2 promoter (mIL2P)

cattttgacacccccataatatTTTTTccagaattaacagtataaattgcattctcttgttcaagagttccctatcactc  
tctttaatcactactcacagtaacctcaactcctgc (SEQ ID NO:1635)

Minimal promoter, miniP  
(DNA)

Agagggtatataatggaagctcgacttccag (SEQ ID NO:1636)

IL-18 activated  
(DNA)

atggccttaccagtgaccgccttgctcctgccgctggccttgetgtctccacgcgcagccagggcgagggtgacttccggaagctg  
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ID NO:1637)  
(amino acids)

MALPVTALLLPLALLLHAARPYFGKLESKLSVIRNLNDQVLFIDQGNRPLFEDMTDSDCR  
DNAPRTIFIISMYKDSQPRGMAVTISVKCEKISTLSCENKIISFKEMNPPDNKDTKSDI  
IFFQRSVPGHDNKMQFESSYEGYFLACEKERDLFKLILKKEDELGDRSIMFTVQNE\*\* (SEQ ID NO:1638)

Foxp3-NFATc1-X3-IL-18

(DNA)

ggcttcattttttccatttactgcagaggcttcattttttccatttactgcagaggcttcattttttccatttactgc  
agaactagtttaggcgtgtacgggtgggaggcctatataagcagagctcgtttagtgaaccgtcagatcgctggagacg  
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NO:1639)

Foxp3-NFATc1-X6-IL-18

(DNA)

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ID NO:1640)

IL-2-NFATc1-X3-IL-18

(DNA)

ggaggaaaaactgtttcatacagaaggcgtggaggaaaaactgtttcatacagaaggcgtggaggaaaaactgtttca  
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a (SEQ ID NO:1641)

IL-2-NFATc1-X6-IL-18

(DNA)

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aggactgataa (SEQ ID NO:1642)

Best antibodies with inducible IL-18 +/- 1XX mutations in ITAMs of CD3-zeta  
MNC2

muMNC2-8-4-1BB-3z-Foxp3-NFAT-IL-18

(DNA)

atggccttaccagtgaccgccttgctcctgcccgtggccttgctgctccacgcgcgcagggccggaggtccagctggag  
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hu25E6-8-28-3z-Foxp3-NFAT-IL-18  
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mu25E6-8-4-1BB-3z1XX-Foxp3-NFAT-IL-18

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(DNA)

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mu25E6-8-28-3z1XX-Foxp3-NFAT-IL-18  
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hu25E6-8-28-3z1XX-Foxp3-NFAT-IL-18

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<b>Construct Name:</b>	<b>pCDH MSCV h20A10-O CAR 41BB</b>
<b>Construct backbone:</b>	<b>pCDH CMV MCS (SBI)</b>
<b>Promoter</b>	<b>MSCV</b>
<b>Leader sequence:</b>	<b>Human CD8 alpha (1- 63)</b>
<b>scFv Name:</b>	<b>Humanized 20A10-O (64 - 807)</b>
<b>Hinge region:</b>	<b>Human CD8 alpha (808 - 942)</b>
<b>Transmembrane Domain:</b>	<b>Human CD8 alpha (943 - 1014)</b>
<b>Costimulatory Domains:</b>	<b>Human 41BB (1015 - 1140) and CD3 zeta (1141 - 1476)</b>





Construct Name:	pCDH MSCV h20A10-N CAR 41BB
Construct backbone:	pCDH CMV MCS (SBI)
Promoter	MSCV
Leader sequence:	Human CD8 alpha (1- 63)
scFv Name:	Humanized 20A10-N (64 - 807)
Hinge region:	Human CD8 alpha (808 - 942)
Transmembrane Domain:	Human CD8 alpha (943 - 1014)
Costimulatory Domains:	Human 41BB (1015 - 1140) and CD3 zeta (1141 - 1476)

[illegible]

CD8 leader sequence

ATGGCCTTACCAGTGACCGCCTTGCTCCTGCCGCTGGCCTTGCTGCTCCACGCCGCCAGGCCG (SEQ ID NO: 1679)

h20A10-N scFV

CAGGTG CAGCTT GGTG AATCT GGC GCGG GACTT GTG AAGCCT GGC GGA TCTCT GAGACT GAGCT GTG CCGCC AGCGCG  
TTCACCTTT TAGCACA TACGCC ATGAGCTGGATCAGACAGGCCCTGGCAAAGGCCTGGAATGGGTGgcgTCTATTGGC  
AGAGCCGGCAGCACCTACTACAGCGACTTGTGTGAAGGGCAGATTACCATCAGCCGGGACAACGCCAAGAACAGCCTG  
TACCTGCAGATGAAC TCCCTGACAGCCGAGGACACCGCCCTGTACTATTGTGCCAGAGGACCCATCTACAACGACTAC  
GACGAGTTCGCC TATTGGGGCCAGGGCACACTGGTTCAGATTTCTAGCGCGGTGGCGGAagcGGAGGCGGTGGCTcc  
GGTGGCGGAGGCagcGAAATTGTGCTGACACAGAGCCCCGCCACACTGTCACTTTCTCCAGGCGAAAGAGCCACACTG  
AGCTGCAAGAGCAGCCAGAGCGTGCTGTACTCCAGCAACCAGAAGAATACTACCTGGCCTGGTATCAGCAGAAGCCCCGGC  
CAAGCTCCTCGGCTGCTGATCTATTGGGGCCAGCACAAAGAGAGAGCGGCATCCCTGCCAGATTTTCTGGCAGCGGCTCT  
GGCACCGATTTTACCCTGACCATAAGCAGCCTGGAACCTGAGGACTTCGCGCGTGTATTACTGCCACCAGTACCTCAGC  
AGCCTGACCTTTTGGCGGAGGCACCAAGGTGGAAATCAAGCGG (SEQ ID NO: 1680)

CD8 hinge region

ACAACAACCCCTGCCCCAGACCTCTTACCCCAGCCCCTACAATTGCCAGCCAGCCTCTGAGCCTGAGGCCCGAGGCT  
TGTAGACCTGCTGCTGGCGGAGCCGTGCACACCAGAGGACTGGATTTTCGCCTGCCAC (SEQ ID NO: 1681)

CD8 transmembrane domain

A C T A C A T T G G C G T T T C T T G A C T T T C G G T T T C G C C T T C G C G C T T A C A C T T A C T C (SEQ  
 ID NO: 1682)

4444

(SEQ ID NO: 1683)



CD3

AGACTGAAGTTCAGCAGGAGCGGCAGACCGCCCGCGCTACAGGCAGCGCCAGCAACAGCTCTATAACCAGCTCAATCTA  
GCACGAAGACAGGAGTACGATGTTTTGGACAAGACACGTCGGCGCGCACCGTCAGATGGGGGCAAGCGCGAGCAAGCAAC  
AAGCCCTCAGGAAGGCGCTGTACAACTGAAGTGCAGAAAGAAGATGGCGGAGCGCTACAGTGCAGATTGGCAAGAAAGGC  
CAGCGCGCGCAGGGGCAAGGGGCACGATGGCCCTTACCAGCGTCTCAGTACAGCCACCAAGGACACCTAGCACCGCCCT  
CACATGCAGCGCGCTGGCGCGCTGGCTGATAA (SEQ ID NO: 1684)

Construct Name:	pCDH MSCV h20A10-C2 CAR 41BB
Construct backbone:	pCDH CMV MCS (SBI)
Promoter	MSCV
Leader sequence:	Human CD8 alpha (1- 63)
scFv Name:	Humanized 20A10-C2 (64 - 807)
Hinge region:	Human CD8 alpha (808 - 942)
Transmembrane Domain:	Human CD8 alpha (943 - 1014)
Costimulatory Domains:	Human 41BB (1015 - 1140) and CD3 zeta (1141 - 1476)

[illegible]

CD8 leader sequence

ATGGCCCTTACCAGTGACCGCCTTGCTCCTGCCGCTGGCCTTGCTGCTCCACGCCGCCAGGCCG (SEQ ID NO: 1686)

## h20A10-C2 scFV

GAGGTGCACGCTGGTTGAATCTGGCGGGCGGACTTGTGAAGCCTGGCGGATCTCTGAGACTGAGCTGTGCCGCCAGCGGG  
TTCACCTTTAGCACATACGCCATGAGCTGGGTCCGACAGGCCCCCTGGAAAAGGCCTTGAATGGGTGGCTCTATCGGG  
AGAGCCGCCAGCACCTACTACAGCGATTCTGTGAAGGGCAGATTACCATCAGCCGGGACAACGCCAAGAACAGCCTG  
TACCTGCAGATGAACTCCCTGAGAGCCGAGGACACCGCCCTGTACTATTGTGCCAGAGGACCCATCTACAACGACTAC  
GACGAGTTCCGCTATTGGGGCCAGGCCACACTCGTCACACTCAGCTCTGGCGGTGGCGGAagcGGAGGCGCTGGCtcc  
GGTGGCGGAGGCagcgacattgtgctgacccagtcctccagcctccttggccgtgtctccaggacagagggccaccatc  
acctgcAAGACCAGCCAGAGCGTGTCTGTACTCCAGCAACCAGAAGAACTACCTGGCctgggtatcagcagaaaccagga  
caacctcctaaactcctgattttacTGGGCCAGCACCAGAGAAAGCgggggtcccagccaggttcagcgggcagtggggtct  
gggacggatttcaacctcacaattaatcctgtggaagctaataatgatactgcaaattattactgtCACCAGTACCTGAGC  
AGCCTCACCTTCGGCGGAGCGACCAAGCTGGAGATCAAACGA (SEQ ID NO: 1687)

CD8 hinge region

ACAAACAACCCCTGCCCCAGACCTCCTACCCCAGCCCTACAATTGCCAGCCAGCCTCTGAGCCTGAGGCCCGAGGCT  
TGTAGACCTGCTGCTGGCGGAGCCGTGCACACCAGAGGACTGGATTTTCGCTGCCAC (SEQ ID NO: 1688)

CD8 transmembrane region

A T C A C A G T G G C A C T T T G G C T T A C T T T T G G C T T C C T T A G G T A C A C C T A C G (SEQ  
ID NO: 1689)



(SEQ ID NO: 1690)

CD3

AGACTGAAGTTCAGCAGGAGCGGCAGACCGCCCCCGCGTACAGGCAGCGCCAGCAACAGCTCTATAACCGAGCTCAATCTA  
GCACGAAGACAGGAGTACGATGTTTTGGACAAGACACGTCGGCGCGCACCGTCAGATGGGGCGAAAGCGGAGAACCAAC  
AAGCCCTCAGGAAGGCGCTGTACAACTGAAGTGCAGAAAGAATAGATGGCGGAGCGCTACAGTGAAGATTGGCAAGAAAGGC  
GAGCGCGCGAGGGGCAAGGGGCGAGGATGGCGCTTACCAGCGTCTCAGTACAGCCACCAAGGACACCTAGGACCGCGCT  
CACATGCAGCGCGTGGCGCGCTCGGTGATAA (SEQ ID NO: 1691)

Construct Name:	pCDH MSCV h20A10-O CAR CD28 1XX
Construct backbone:	pCDH CMV MCS (SBI)
Promoter	MSCV
Leader sequence:	Human CD8 alpha (1- 63)
scFv Name:	Humanized 20A10-O (64 - 807)
Hinge region:	Human CD8 alpha (808 - 942)
Transmembrane Domain:	Human CD8 alpha (943 - 1014)
Costimulatory Domains:	Human CD28 (1015 - 1137) and CD3 zeta 1XX (1138 - 1473)

ATGGCCCTTACCAGTGACCGCCTTGTCTCTGCCGCTGGCCTTGTCTGCTCCACGCCGCCAGGCCGAGGTCACAGCTGGTT  
GAATCTGGCGCGGCACTTGTGAAGCCTGGCGGATCTCTGAGACTGAGCTGTGCCGCCAGCGGCTTCACCTTTAGCACA  
TACGCCATCAGCTGGGTCCGACAGGCCCTGGAAAAGGCCCTTCAATGGGTTGCCTCTATCGGCAGAGCCGGCAGCACC  
TACTACAGCGATTCTGTGAAGGGCAGATTACCATCAGCCGGGACAACGCCAAGAACAGCCTGTACCTGCAGATGAAG  
TCCCTGACAGCCGAGGACACCGCCGTGTACTATTGTGCCAGAGGACCCATCTACAACGACTACGACGAGTTCGCCCTAT  
TGGGGCCAGGCCACACTGGTTCAGAGTCAGCTCTGGCGGTGGCGGAagcGGAGGCGGTGGGctcGGTGGCGGAGGGcagc  
GACATCGTGATGACACAGAGCCCTGATAGCCTGGCCGTGTCTCTGGGAGAGAGAGCCACCATCAACTGCAAGAGCAGC  
CAGACCGTGCTGTACTCCAGCAACCAGAAGAACTACCTGGCCTGGTATCAGCAGAAGCCCGGCCAGCCTCCTTAAGCTG  
CTGATCTACTGGGCCAGCACCAGAGAAAGCGGCGTGCCCGATAGATTTTCTGGCAGCGGCTCTGGCACCGACTTCACC  
CTGACAATTAGCTCCCTGCAAGCCGAGGATGTGGCCGTGTACTACTGTCCACAGTACCTGAGCAGCCTGACCTTTGGC  
GGCGCAACAAAGGTGGAAATCAAGcgaACAACAACCCCTGCCCCCAGACCTCCTACCCCAGCCCCCTACAATTGCCAGC  
CAGCCTCTGAGCCTCAGCCCCGAGGCTTGTAGACCTGCTGCTGGCGGAGCCGTGCACACCAGAGGACTGGATTTGCC  
TGGGACATCAGATCTGGGCGCCTTGGCGTGAAGCTGTGGGTCCTCTGGCTGTACCTGATCAAGCTTACTG  
AGCAATAGAGGACAGGCTTCTGACAGTCACTACATGAACATGAGCCAGAGAGAGCGGCCAGCAAGAAAG  
TACAGCGCTATGCGGAGCAGCGGACTGGGAGCTAAGGCTCAGAGTCAAGTTCACAGGAGCCAGACAGCCCC  
GCTTACAAGCAGGCCAGAACCACTCTATAACGAGCTCAATCTAGGACGAAGAGAGGAGTACCATGTTTGGACAAG  
ACAGCTGGCGCGGACCCCTCAGATGCCCCGAAACCCGAGAAGGAAGAACCCTCAGGAAGCCCTCTTCAATGAATGCAG  
AAAGATAAGATGGCGGACGCCCTTCAGTCAGATTGGGATGAAAGGCCACCGCCCGAGGGGCAAGGGGCACCATGGCCCT  
TTCAGGGGTCTCAGTACAGCCACCAAGCACACCTTCCAGCCCTTTCACATGCAGGCCCTGGCCCTTCGCTGATAA  
(SEQ ID NO:1692)

CD8 leader sequence  
ATGGCCCTTACCAGTGACCGCCTTGCTCCTGCCGCTGGCCTTGCTGCTCCACGCCGCCAGGCCG (SEQ ID NO:  
1693)

h20A10-0 scEV

GAGGTGCACGCTGGTTGAATCTGGCGGCGGACCTTGTGAAGCCTGGCGGATCTCTGAGACTGAGCTGTGCCGCCAGCGGC  
TTCACCTTTAGCACATACGCCATGAGCTGGGTCCGACAGGCCCTTGGAAAAGGCCTTGAATGGGTGGCTCTATCGGC  
AGAGCCGCCACACCTACTACACCGATTCTGTGAAGGGCAGATTACCATCAGCCGGGACAACGCCAAGAACAGCCTG  
TACCTGCAGATGAACCTCCCTGCAGCCGAGGACACCGCCCTGTACTATTGTGCCAGAGGACCCATCTACAACGACTAC  
GACGAGTTCGCCTATTGGGGGCCAGGGCACACTGGTCACAGTCAGCTCTGGCGGTGGCGGAagcGGAGGCGGTGGCTcc  
GGTGGCGGAGGCagcGACATCGTGATGACACAGAGCCCTGATAGCCTGGCCGTGTCTCTGGGAGAGAGAGCCACCATC  
AACTGCAAGACCAGCCAGAGCGTGTCTGTACTCCAGCAACCAGAAGAACTACCTGGCCTGGTATCAGCAGAAGCCCGGC  
CAGCCTTCCTAAGCTGCTGATCTACTGGGCCAGCACCAGAGAAAGCGGCGTGGCCGATAGATTTTCTGGCAGCGGCTCT  
GGCACCAGACTTCACCCTGACAATTAGCTCCCTGCAGGCCGAGGATGTGGCCGTGTACTACTGTACCAGTACCTGAGC  
AGCCTGACCTTTGGCGGCGGAACAAAGGTGGAAATCAAGcga (SEQ ID NO: 1694)

CD8 hinge region

ACAACAACCCCTGCCCCAGACCTCCTACCCCAGCCCCTACAATTGCCAGCCAGCCCTCTGAGCCTGAGGCCCGAGGCCTG  
TGTAGACCTGCTGCTGGCGGAGCCGTGCACACCAGAGGACTGGATTTTCGCCTGCCAC (SEQ ID NO: 1695)

CD8 transmembrane region

AATGACAACTTGGGGCGGTTCCTGGGGTTCACTGTTAGGAGTTTCCTCCTTAPCATGCCTTACATG (SEQ ID NO: 1696)



(SEQ ID NO: 1697)

CD3 1XX

AGACTGAAGTTCAGCAGGAGCGGCAGACCGCCCGCGCTACAGGCAGCGCCAGCAACAGCTCTATAACCAGCTCAATCTA  
GCACGAAGACAGGAGTACCATGTTTTGGACAAGACACGTCGGCGCGCACCGTCAGATGGGGGCAAGCGCGAGCAAGCAAC  
AAGCCCTCAGGAAGGCGCTGTCAATGAAGTGCAGAAAGAAGATGGCGGAGCGCTTCAGTCAGATTGGCAAGAAAGGC  
CAGCGCGCGCAGGGGCAAGGGGCACGATGGCCCTTCCAGCGTCTCAGTACAGCCACCAAGGACACCTTCGACCGCGCT  
CACATGCAGCGCGTGGCGCGCTCGGTGATAA (SEQ ID NO: 1698)



AGACTGAAGTTCAGCAGGAGCGGCAGACCCCCCGCGTACAGGCAGCGCCAGAACAGCTCTATAACCGAGCTCAATCTA  
GCACGAAGACAGGAGTACGATGTTTTGGACAAGACACGTCGGCGCGCACCGTCAGATGGGGGCAAGCGCGAGAACGAAC  
AAGCCCTCAGGAAGGCGCTGTCAATGAAGTGCAGAAAGAAGATGGCGGAGCGCTTCAATGAGATTGGCAAGAAAGGC  
GAGCGCGCGAGGGGCAAGGGGCAGGATGGCCCTTCCAGCGTCTCAGTACAGCCACCAAGGACACCTTCCACCGCGCT  
CACATGCAGCGCGTGGCGCGCTGGCTGATAA (SEQ ID NO: 1705)



Construct Name:	pCDH MSCV h20A10-C2 CAR CD28 1XX
Construct backbone:	pCDH CMV MCS (SBI)
Promoter	MSCV
Leader sequence:	Human CD8 alpha (1- 63)
scFv Name:	Humanized 20A10-C2 (64 - 807)
Hinge region:	Human CD8 alpha (808 - 942)
Transmembrane Domain:	Human CD8 alpha (943 - 1014)
Costimulatory Domains:	Human CD28 (1015 - 1137) and CD3 zeta 1XX (1138 - 1473)

ATGCGCCTTACCAGTGACCGCCCTTGCTCCTGCCGCTGGCCTTGCTGCTCCACGCCGCCAGGCCCGAGGTCACAGCTGGTT  
 GAATCTGGCGCGCGACTTGTGAAGCCTGGCGGATCTCTGAGACTGAGCTGTGCCGCCAGCGGCTTCACCTTTAGCACA  
 TACGCCATCAGCTGGGTCCGACAGGCCCTTGGAAAAGGCCCTTCAATGGGTTGCCTCTATCGGCAGAGCCGGCAGCACC  
 TACTACAGCGATTCTGTGAAGGGCAGATTACCATCAGCCGGGACAACGCCAAGAACAGCCTGTACCTGCAGATGAAG  
 TCCCTGACAGCCGAGGACACCGCCGTGTACTATTGTGCCAGAGGACCCATCTACAACGACTACGACGAGTTCGCCCTAT  
 TGGGGCCAGGCCACACTGGTCAAGTCAGCTCTGGCGGTGGCGGAagcGGAGGCGGTGGCTcggGTGGCGGAGGGcagc  
 gacattgtggtgacccagtgctccagcctccttgggcgtgtctccaggacagagggccaccatcacctgcAAGAGCAGC  
 CAGACCGTGCTGTACTCCAGCAACCAGAAGAACTACCTGGCCTggtatcagcagaaaaccaggacaacctcctaaactc  
 ctgattttacTGGGCCAGCACCAGAGAAAGCGggggtcccagccagggttcagcggcagtggggtctgggaccgatttcacc  
 ctcaacaattaatcctgtggaagctaataatgatactgcaattattactgtCACCAGTACCTGAGCAGCCTGACCTTCGGG  
 GGAGGCACCAAGGTGGACATCAAACGAACAACAACCCCTGCCCCAGACCTCCTACCCCAGCCCCCTACAATTGCCCAGC  
 CAGCCTCTGAGCCTCAGCCCCGAGGCTTGTAGACCTGCTGCTGGCGGAGCCGTGCACACCAGAGGACTGGATTTCGCC  
 TGGCACA CTACATCTGGGCGCCCTTGGCGTGGACCTGTGGGTGGCTCTGGCTGTCACTGGCTATCAGCCCTACCTG  
 CTGCAATAAGGATCAGGCTCTGCAAGTCACTACATGAACATGAGCGTAGAAGACGGGCCAGCAGAAAGCA  
 TACAGCCCTATCTGGGACAGCGGACTGGGAGCTAAGGCTCAGAGTCAAGTTCACAGGACGCGACAGCGCCCC  
 CGCTACAAGCAGGGCCAGAACCAGCTCTATAACGAGCTCAATCTAGGACGAAGAGAGGAGTACCATGTTTGGACAAG  
 ACAGCTGGCGCGGACCCCTCAGATGCCCCGAAACCCGAGAAGGAAGAACCCTCAGGAAGCCCTCTTCAATGAACGCAAG  
 AAAGATAAGATGGCGGACGCCCTTCAGTCAGATTGGGATGAAAAGGCCAGCGCCCGAGGGGCAAGGGGCACCATGGCCCT  
 TCCAGGGTCTCAGTACAGCCACCAAGCACACCTTCCAGCCCCCTTCACATGCAGGCCCTGCCCCCTGGCTGATAA  
 (SEQ ID NO:1706)

CD8 leader sequence  
ATGGCCCTTACCAGTGACCGCCTTGCTCCTGCCGCTGGCCTTGCTGCTCCACGCCGCCAGGCCG (SEQ ID NO:  
1707)

## h20A10-C2 scFV

GAGGTGCAGCTGGTTGAATCTGGCGCGCGGACTTGTGAAGCCTGGCGGATCTCTGAGACTGAGCTGTGCCGCCAGCGGC  
TTCACCTTTAGCACATACGCCATGAGCTGGGTCCGACAGGCCCTTGGAAAAGGCCTTGAATGGGTGGCTCTATCGGC  
AGAGCCGCCAGCACCTACTACAGCGATTCTCTGAAGGGCAGATTACCATCAGCCGGGACAACGCCAAGAACAGCCTG  
TACCTGCAGATGAACTCCCTGAGAGCCGAGGACACCGCCGTGTACTATTGTGCCAGAGGACCCATCTACAACGACTAC  
GACGAGTTCGCCCTATTGGGGCCAGGGCACACTGGTCACAGTCAGCTCTGGCGGTGGCGGAagcGGAGGCGGTGGctcc  
GGTGGCGGAGGCagcgacattgtgctgacccagttctccagcctccttgccgtgtctccaggacagagggccaccatc  
acctgcAAGACCAGCCAGAGCGTGTCTGTACTCCAGCAACCAGAAGAACTACCTGGCctgggtatcagcagaaaccagga  
caacctcctaaactcctgatttacTGGGCCAGCACCAGAGAAAGCgggggtccagccaggttcagcggcagtggtgt  
gggaccgatctcaccctcacattaatcctgtggaagctaatgatactgcaaattattactgtCACCAGTACCTGAGC  
AGCCTGACCTTCGCCGGAGGGACCAAGGTGGAGATCAAACGA (SEQ ID NO: 1708)

## CD8 hinge region

ACAACAACCCCTGCCCCAGACCTCCTACCCCAGCCCCTACAATTGCCAGCCAGCCTCTGAGCCTGAGGCCCGAGGGCT  
TGTAGACCTGCTGCTGGCGGAGCCGTGCACACCAGAGGACTGGATTTGCCTGCCAC (SEQ ID NO: 1709)

## CD8 Transmembrane region

A C C A A C T G G T T T C T G T T G A A T T G G G T T T C C A T T C A C T T T A C A T T T A T T T (SEQ  
 ID NO: 1710)

(SEQ ID NO: 1711)

(SEQ ID NO: 1712)

[illegible]

C2\_scFV

GAGGTCGACGTCGGTCGACTCTGGGGGAGGCCTGGTCAAGCCTGGGGGGTCCCTGAGACTCTCCTGTGTCAGCCCTCTGGA  
TTCACCTTCACTGGCTATGCCATGAGCTGGGTCCGCCAGGCCTCCAGGGAAGGGGCTGGAGTGGGTCTCAACCATTAGT  
AGTGGCGGAACCTACATATACTACCCCGACTCAGTGAAGGGGCGATTACCATCTCCAGAGACAACGCCAAGAAGCTCA





## CD3 1XX

AGASTCAAGTTCAGCAGGAGCCGAGACGCCCCCGCGGTACAAGCAGCCCCAGAACCCAGCTCTATAACCCAGCTCAATCTA  
 SEACGAACAGAGGAGTACGATGTTTGGACAAGAGACGTCGCCCCGACCCCTGAGATGGGGGAAAAGCCGACAAGCAAG  
 AACCCCTCAGCAAGGCCGTGTTCAATGAAGTCCAGAAAGATAAGATGGCCGAGCCCTTCAGTCACATTGGCAACAAAGGC  
 GAGCGCCCGAGGGGCAAGGGGCACGATGGCCCTTCCAGGGTCTCAGTACAGCCACCAAGGACACCTTCGACGCCCTT  
 CACATGCAGGCCCTCGCCCCCTCGGTGATAA (SEQ ID NO: 1732)

Mouse B12 light chain variable *framework 2 (FR2)* sequence:  
 (DNA)

tggtatcagcagaaaccaggacagccacccaaactcctcatctat (SEQ ID NO: 1733)

(amino acids)

WYQQKPGQPPKLLIY (SEQ ID NO: 1734)

Mouse B12 light chain variable *complementarity determining regions 2 (CDR2)* sequence:

(DNA)

cttgcattccaccctagattct (SEQ ID NO: 1735)

(amino acids)

LASTLDS (SEQ ID NO: 1736)

Mouse B12 light chain variable *framework 3 (FR3)* sequence:  
 (DNA)

ggggtccctgccaggttcagtggtcagtggttctaggacagacttcaccctcaccattgatcctgtggaggctgatgat

gctgcaacctattactgt (SEQ ID NO: 1737)

(amino acids)

GVPARFSGSGSRTDFTLTIDPVEADDAATYYC (SEQ ID NO: 1738)

Mouse B12 light chain variable *complementarity determining regions 3 (CDR3)* sequence:

(DNA)

cagcaaaataatgaggatcctccgacg (SEQ ID NO: 1739)

(amino acids)

QQNNEDPPT (SEQ ID NO: 1740)

Mouse B12 light chain variable *framework 4 (FR4)* sequence:  
 (DNA)

ttcgggtggaggcaccaagctggaaatcaagg (SEQ ID NO: 1741)

(amino acids)

FGGGTKLEIK (SEQ ID NO: 1742)

epitope to which NME1 and NME7<sub>AB</sub> bind part of the 10 membrane proximal amino acids

**PFPSAQSGA** (SEQ ID NO: 1743)

SNIKFRPGSVV (SEQ ID NO: 1744)

ASRYNLT (SEQ ID NO: 1745)

Fragment of PSMGFR

GTINVHDTVET (SEQ ID NO: 1746)

Fragment of PSMGFR

FPFS (SEQ ID NO: 1747)

Fragment of PSMGFR

SNIKFRPGSVVVQLTLAFRE (SEQ ID NO: 1748)

Fragment of PSMGFR  
QFNQYKTEA (SEQ ID NO:1749)

Fragment of PSMGFR  
VQLTLAFRE (SEQ ID NO:1750)

Fragment of PSMGFR  
SVSDV (SEQ ID NO:1751)

**[00820]** IL18 sequence

**[00821]** (DNA)

**[00822]** Atagaagacaccgggaccgatccagcctcgagagacccaatgctagccaccatggccttaccagtgaacgccttgctcctgcccgtggccttgctgctccacgcgcgcagggcgtacttcggcaagctggaaagcaagctgagcgtgatccggaacctgaacgaccaggtgctgttcacatcgatcagggcaacagacccctgttcgaggacatgaccgacagc gactgcagagacaacgcccctcgaccatcttcacatcagcatgtacaaggacagccagcctagaggcatggccgtgaccatctctgtgaagtgcgagaagatcagcaccctgagctgcgagaacaagatcatcagcttcaaagagatgaaccgcccggacaacatcaaggacaccaagagcgcacatcatattcttcagcggagcgtgcccggccacgacaacaagatgcagtttgagagcagcagctacgagggctacttccctggcctgcgagaaaagagcgggacctgttcaagctgatccctgaa gaaagaggacgaactgggcgaccgcagcatcatgttcaccgtgcagaacgaggactgataaaagcttggcaatccgg tactgttggttaaagccacca (SEQ ID NO:1752)

**[00823]** (amino acids)

IEDTGTDPASRDPLATMALPVTALLLPLALLLHAARPYFGKLESKLSVIRNLNDQVLFI DQGNRPLFEDMTDSDCR  
DNAPRTIFIISMYKDSQPRGMAVTISVKCEKISTLSCENKIISFKEMNPPDNIKDTKSDIIFQ RSVPGHDNKMQFE  
SSSYEGYFLACEKERDLFKLILKKEDELGDRSIMFTVQNE D\*\*KLGNPVLLVKPP (SEQ ID NO:1753)

primer

5'- agggagaccaagctggctagttaagcttggatggccttaccagtgaccgccttgc-3' (SEQ ID NO:1754)

primer

5'- taggccagagaaatgttctggcattatcagcgagggggcagggcctgc-3' (SEQ ID NO:1755).

5'- tgccagaacatttctcttg-3' (SEQ ID NO:1756)

5'- acagtcgaggctgatcagcgggttttaaacttatcagtcctcgcttctgcacgg-3' (SEQ ID NO: 1757)

5'-atgcaggccctgccccctcgctgataagtttaaactgccagaacatttctctggcctaac-3' (SEQ ID NO:1758)

5'- accggagcgatcgagatccttcgcggccgcttatcagtcctcgcttctgcacgggtgaac-3' (SEQ ID NO:1759)

5'- attgcactagttgaaagacccacctgtagg-3' (SED ID NO:1760)

5'- aatgctctagaatacgggtatccagg-3' (SEQ ID NO:1761)

5' atagcgaattcgtaccgagggccaccatgg-3' (SEQ ID NO:1762)

5'- taggcctcccaccgtacacgcctaggtaccacgccttctgtatg-3' (SEQ ID NO:1763)

5'- taggcctcccaccgtacacgcctaggtacctctgcagtaaattgg-3' (SEQ ID NO:1764)

5'- taaggccatggtggctagc-3' (SEQ ID NO:1765)

5'- aataagttttaaactgccagaacattttctctgg-3' (SEQ ID NO:1766)

5'- atatagcggccgcttatcagtcctcgttctgcacgg-3' (SEQ ID NO:1767)

**[00824]** 6x FoxP3NFAT mCMV

**[00825]** (DNA)

**[00826]** Agaacatttctctggcctaactggccggtaccgggttcattttttccatttactgcagagggttcatt  
ttttccatttactgcagagggttcattttttccatttactgcagagggttcattttttccatttactgcagagggtt  
cattttttccatttactgcagagggttcattttttccatttactgcagaaactagttaggcgtgtacgggtgggaggcc  
tatataagcagagctcgtttagtgaaccgtcagatcgctggagacgccatccacgctgtttgacctccatagaag  
acaccggggaccgatccagcctcgagagacccaatgctagccaccatgg (SEQ ID NO:1768)

[00827] (amino acids)

**[00828]** RTFLWPNPVPASFFPPTAEASFFPPTAEASFFPPTAEASFFPPTAEASFFPPTAEASFFPPTAEASFFPPTAELVR  
RVRWEAYISRARLVNRQIAWRRHPRCFDLHRRHRDRSSLERPNAHHG (SEQ ID NO:1769)

**[00829]** 6x FoxP3NFAT mIL2P

**[00830]** (DNA)

**[00831]** Agaacatttctctggcctaactggccggtaccggcttcattttttccatttactgcagaggcttcatt  
ttttccatttactgcagaggcttcattttttccatttactgcagaggcttcattttttccatttactgcagaggctt  
cattttttccatttactgcagaggcttcattttttccatttactgcagaactagtcattttgacacccccataatat  
ttttccagaattaacagtatataaattgcatctcttggtcaagaggttccctatcactctctttaatcaactactcacagt  
aacctcaactcctgctcgagagacccaatgctagccaccatgg (SEQ ID NO:1770)

[00832] (amino acids)

**[00833]** RTFLWPNWPVPASFFPFTAEASFFPFTAEASFFPFTAEASFFPFTAEASFFPFTAEASFFPFTAEASFFPFTAEALVI  
LTPP\*YFSRINSINCISCSRVPHYSL\*SLLTSTPASRDPLATMX (SEQ ID NO:1771)

**[00834]** 6x FoxP3NFAT miniP

**[00835]** (DNA)

**[00836]** Agaacatttctctggcctaactggccggtaccggcttcattttttccatttactgcagaggcttcatt  
ttttccatttactgcagaggcttcattttttccatttactgcagaggcttcattttttccatttactgcagaggctt  
cattttttccatttactgcagaggcttcattttttccatttactgcagaactagtagagggtatataatggaagctc  
gacttccagctcgagagaccaatgctagccaccatgg (SEQ ID NO:1772)

[00837] (amino acids)

**[00838]** RTFLWPNWPVPASFFPFTAEASFFPFTAEASFFPFTAEASFFPFTAEASFFPFTAEASFFPFTAEALVE  
GI\*WKLDFOLEPNASHHG (SEQ ID NO:1773)

**[00839]** 6x IL2NFAT mCMV

**[00840]** (DNA)



**[00841]** Agaacatttctctggcctaactggccggtaccggaggaaaaactgtttcatacagaaggcgtggagga  
 aaaactgtttcatacagaaggcgtggaggaaaaaactgtttcatacagaaggcgtggaggaaaaaactgtttcatacag  
 aaggcgtggaggaaaaaactgtttcatacagaaggcgtggaggaaaaaactgtttcatacagaaggcgtactagttagg  
 cgtgtacggtgggaggcctatataagcagagctcgttttagtgaaccgtcagatcgctggagacgccatccacgtg  
 ttttgacctccatagaagacaccgggaccgatccagcctcgagagacccaatgctagccaccatgg (SEQ ID  
 NO:1774)

**[00842]** (amino acids)

**[00843]** RTFLWPNWPVPEEKLFHTEGVEEKLFHTEGVEEKLFHTEGVEEKLFHTEGVEEKLFHTEGVEEKLFHT  
 EGVLVRRVRWEAYISRARLVNRQIAWRRHPRCFDLHRRHRDRSSLERPNSHHG (SEQ ID NO:1775)

**[00844]** 6x IL2NFAT mIL2P

**[00845]** (DNA)

**[00846]** Agaacatttctctggcctaactggccggtaccggaggaaaaactgtttcatacagaaggcgtggagga  
 aaaactgtttcatacagaaggcgtggaggaaaaaactgtttcatacagaaggcgtggaggaaaaaactgtttcatacag  
 aaggcgtggaggaaaaaactgtttcatacagaaggcgtggaggaaaaaactgtttcatacagaaggcgtactagtcatt  
 ttgacacccccataatatTTTTTccagaattaacagtataaattgcatctcttgttcaagagttccctatcactctct  
 ttaatcactactcacagtaacctcaactcctgcctcgagagacccaatgctagccaccatgg (SEQ ID  
 NO:1776)

**[00847]** (amino acids)

**[00848]** RTFLWPNWPVPEEKLFHTEGVEEKLFHTEGVEEKLFHTEGVEEKLFHTEGVEEKLFHTEGVEEKLFHT  
 EGVLVILTPP\*YFSRINSINCISCSRVPHYSL\*SLLTVTSTPASRDPMLATMX (SEQ ID NO:1777)

**[00849]** 6x IL2NFAT miniP

**[00850]** (DNA)

**[00851]** Agaacatttctctggcctaactggccggtaccggaggaaaaactgtttcatacagaaggcgtggagga  
 aaaactgtttcatacagaaggcgtggaggaaaaaactgtttcatacagaaggcgtggaggaaaaaactgtttcatacag  
 aaggcgtggaggaaaaaactgtttcatacagaaggcgtggaggaaaaaactgtttcatacagaaggcgtactagtagag  
 ggtatataatggaagctcgacttccagctcgagagacccaatgctagccaccatgg (SEQ ID NO:1778)

**[00852]** (amino acids)

**[00853]** RTFLWPNWPVPEEKLFHTEGVEEKLFHTEGVEEKLFHTEGVEEKLFHTEGVEEKLFHTEGVEEKLFHT  
 EGVLVEGI\*WKLDQLERPNASHHG (SEQ ID NO:1779)

#### **Human NME1**

(DNA)

atggccaactgtgagcgtaccttcattgcatcaaaccagatgggggtccagcgggggtcttgtgggagagattatcaa  
 gcgttttgagcagaaaaggattccgccttggtggtctgaaattcatgcaagcttcgaagatcttctcaaggaaact  
 acgttgacctgaaggaccgtccattctttgcccgtggtgaaatacatgcactcagggccggttagttgccatgggtc  
 tgggaggggctgaatgtggtgaagacggggccagtcgtgctcggggagaccaaccctgcagactccaagcctgggac  
 catcgtggagacttctgcatacaagttggcaggaacattatacatggcagtgattctgtggagagtgagagagaagg  
 agatcggccttggtgtttcacccctgaggaactggttagattacagagctgtgctcagaactggatctatgaatga  
 (SEQ ID NO:1780)

(amino acids)

MANCERTFIAIKPDGVQRGLVGEIIKRFEQKGFRLVGLKFMQASEDLLKEHYVDLKDRPFFAGLVKYMHS GPVVAMV  
WEGLNVVKTGRVMLGETNPADSKPGTIRGDFCIQVGRNIIHGSDSVESA EKEIGLWFHPEELVDYTS CAQNWIYE-  
(SEQ ID NO:1781)

#### Human NME7

(DNA)

atgaatcatagtgaaagattcgttttcattgcagagtggtatgatccaaatgcttcacttcttcgacggttatgagct  
tttattttaccaggggatggatctgttgaaatgcatgatgtaaagaatcatcgaccccttttaagcggaccaa  
atgataacctgcacttggaagatttatttataggcaacaaagtgaatgtctttctcgacaactgggtattaattgac  
tatggggatcaatatacagctcgccagctgggcagtaggaaagaaaaaacgctagccctaattaaaccagatgcaat  
atcaaaggctggagaaataattgaaataataacaaagctggatttactataaccaaaactcaaaatgatgatgcttt  
caaggaaagaagcatttgattttcatgtagatcaccagtcagagccctttttcaatgagctgatccagtttattaca  
actggtcctattattgccatggagattttaagagatgatgctatatgtgaatggaaaagactgctgggacctgcaaa  
ctctggagtggcacgcacagatgcttctgaaagcattagagccctctttggaacagatggcataaagaaatgcagcgc  
atggccctgattcttttgcctctgcggccagagaaatggagttgtttttccttcaagtggaggttgtgggcccggca  
aacactgctaaatttactaattgtacctgttgcatgttaaaccatgctgtcagtgaggactgttgggaaagat  
cctgatggctatccgagatgcagggtttgaaatctcagctatgcagatgttcaatatggatcgggttaattgttgagg  
aattctatgaagtttataaaggagtagtgaccgaatatcatgacatgggtgacagaaatgtattctggcccttgtgta  
gcaatggagattcaacagaataatgctacaaagacatttcgagaattttgtggacctgctgatcctgaaattgcccg  
gcatttacgccctggaactctcagagcaatctttggtaaaactaagatccagaatgctgttcactgtactgatctgc  
cagaggatggcctattagaggttcaatacttcttcaagatcttggataattag (SEQ ID NO:1782)

#### [00854] (amino acids)

MNHSERFVFI AEWYDPNASLLRRYELLFY PGDGSVEMHDVKNHRTFLKRTKYDNLHLEDLFI GNKVNVS RQLVLID  
YGDQYTARQLGSRKEKTLALIKPDAISKAGEIIIEIINKAGFTITKLKMMMLSRKEALDFHVDHQSRPFFNELIQFIT  
TGPIIAMEILRDDAICEWKRL LGPANSGVARTDASESIRALFGTDGIRNAAHGPDSFASAAREMELFFPSSGGCGPA  
NTAKFTNCTCCIVKPHAVSEGLLGKILMAIRDAGFEISAMQMFNMDRVNVVEEFYEVYKGVVTEYHDMVTEMYSGPCV  
AMEIQQNNATKTFREFCGPADPEIARHLRPGTLRAIFGKTKIQNAVHCTDLPEDGLLEVQYFFKILDN- (SEQ ID  
NO:1783)

**[00855]** Those skilled in the art will recognize, or be able to ascertain using no more than routine experimentation, many equivalents to the specific embodiments of the invention specifically described herein.

What is claimed is:

1. An antibody, or fragment thereof, for the diagnosis, treatment or prevention of cancers wherein the antibody specifically binds to the PSMGFR peptide (SEQ ID NO:2) or a fragment thereof of the peptide.
2. The antibody, or fragment thereof of claim 1, which binds to the N-10 peptide (SEQ ID NO:3), N-19 peptide (SEQ ID NO:4), N-23 peptide (SEQ ID NO:5), N-26 peptide (SEQ ID NO:6), N-30 peptide (SEQ ID NO:7), N-10/C-5 peptide (SEQ ID NO:8), N-19/C-5 peptide (SEQ ID NO:9), or C-5 peptide (SEQ ID NO:825).
3. The antibody, or fragment thereof, of claim 1, which interacts with a peptide comprising conformational epitope SVSDV (SEQ ID NO:1751) and FPSA (SEQ ID NO:1747) within N-26 sequence ISDVSVDVFPFSAQSGA (SEQ ID NO:6), wherein mutation or deletion of FPFS (SEQ ID NO:1747) destroys binding of the antibody or fragment thereof to the N-26 peptide.
4. The antibody, or fragment thereof, of claim 1, which interacts with a peptide comprising conformational epitope ASRYNLT (SEQ ID NO:1745), SVSDV (SEQ ID NO:1751), and FPSA (SEQ ID NO:1747) within the N-19 sequence ASRYNLT ISDVSVDVFPFSAQSGA (SEQ ID NO:4), wherein mutation or deletion of ASRYNLT (SEQ ID NO:1745) destroys binding of the antibody or fragment thereof to the N-26 peptide.
5. The antibody, or fragment thereof, of claim 1, which does not bind to the C-10 peptide (SEQ ID NO:825).
6. The antibody, or fragment thereof of claim 5, which binds to the N-10 peptide (SEQ ID NO:3), but not to the C-10 peptide (SEQ ID NO:825).
7. The antibody, or fragment thereof, of claim 1, which inhibits interaction between NME7<sub>AB</sub> and MUC1\*.
8. The antibody, or fragment thereof, of claim 1, which inhibits interaction between NME7<sub>AB</sub> and PSMGFR peptide (SEQ ID NO:2).

9. The antibody, or fragment thereof, of claim 2, which inhibits interaction between NME7<sub>AB</sub> and N-10 peptide (SEQ ID NO:3), N-19 peptide (SEQ ID NO:4), N-23 peptide (SEQ ID NO:5), N-26 peptide (SEQ ID NO:6), N-30 peptide (SEQ ID NO:7), N-10/C-5 peptide (SEQ ID NO:8), N-19/C-5 peptide (SEQ ID NO:9), or C-5 peptide (SEQ ID NO:825).
10. An antibody, or fragment thereof, of claim 1, wherein the antibody recognizes a MUC1 transmembrane enzymatic cleavage product.
11. The antibody, or fragment thereof, of claim 10, wherein the cleavage enzyme is MMP14 or MMP9 or a catalytically active fragment thereof of the enzyme.
12. The antibody, or fragment thereof, of claim 1, which binds to PSMGFR (SEQ ID NO:2) or fragment thereof in which presence of an amino acid sequence within PSMGFR (SEQ ID NO:2) induces binding of the antibody to the PSMGFR.
13. The antibody of claim 12, wherein the amino acid sequence of the binding conformationally inducing peptide is present in N-10 peptide (SEQ ID NO:3).
14. The antibody, or fragment thereof, of claim 12, which does not bind to a linear form of the binding conformationally inducing peptide sequence wherein the linear form of the peptide is a denatured form.
15. The antibody, or fragment thereof, of claim 14, wherein the binding conformationally inducing peptide sequence is in the N-26 peptide sequence ISDVS<sub>V</sub>SDVPFPFSAQSGA (SEQ ID NO:6), wherein mutation or deletion of FPFS (SEQ ID NO:1747) destroys binding of the antibody or fragment thereof to the N-26 peptide.
16. The antibody, or fragment thereof, of claim 14, wherein the binding conformationally inducing peptide sequence is located within the N-19 sequence ASRYNLTISDVS<sub>V</sub>SDVPFPFSAQSGA (SEQ ID NO:4), wherein mutation or deletion of ASRYNLT (SEQ ID NO:1745) destroys binding of the antibody or fragment thereof to the N-19 peptide.
17. An antibody, or fragment thereof, for the diagnosis, treatment or prevention of cancers wherein a binding inducing peptide sequence is located within the N-26 sequence ISDVS<sub>V</sub>SDVPFPFSAQSGA

(SEQ ID NO:6), wherein mutation or deletion within FPFS (SEQ ID NO:1747) destroys binding of the antibody or fragment thereof to PSMGFR.

18. The antibody, or fragment thereof of claim 17, wherein

heavy chain CDR1 comprises consensus sequence at least 90% identical to sequence: F or I at position 1, T at position 2, F at position 3, S at position 4, T, G, or R at position 5, Y at position 6, A, G or T at position 7, M at position 8 and S at position 9;

heavy Chain CDR2 comprises consensus sequence at least 90% identical to sequence: T at position 1, I or S at position 2, I or S at position 3, G or R at position 5, G or A at position 6, T or I at position 9, Y at position 10, Y at position 11, P or S at position 12 and DSVKG for positions 13-17;

heavy chain CDR3 comprises consensus sequence at least 90% identical to sequence: \_ G, L, or N at position 2, G or T at position 4, Y at position 7, D or E at position 12, A at position 14, and Y at position 15;

light chain CDR1 comprises consensus sequence at least 90% identical to sequence: K or R at position 1, A or S at position 2, S at position 3, K or Q at position 4, S at position 5, L or V at position 6, L at position 7, T or S at position 10, Y at position 15, and I, L or M at position 16;

light Chain CDR2 comprises consensus sequence at least 90% identical to sequence: L or W, or S at position 1, A or T at position 2, S at position 3, N or T at position 4, L or R at position 5, E or A at position 6, and S at position 7; and

light chain CDR3 comprises consensus sequence at least 90% identical to sequence: Q at position 1, H or Q at position 2, S, Q or R at position 3, R, S or Y at position 4, E, L, or S at position 5, L or S at position 6, P or S at position 7, F or L at position 8 and T at position 9.

19. An antibody, or fragment thereof, for the diagnosis, treatment or prevention of cancers wherein binding conformationally inducing peptide is within the N-26 sequence ISDVSVSDVFPFSAQSGA (SEQ ID NO:6), wherein mutation or deletion within FPFS (SEQ ID NO:1747), SVSDV (SEQ ID NO:1751), or ASRYNLT (SEQ ID NO:1745) destroys binding of the antibody or fragment thereof to PSMGFR.

20. The antibody, or fragment thereof of claim 17, wherein  
wherein

heavy chain CDR1 comprises consensus sequence at least 90% identical to sequence: F or I at position 1, T or A at position 2, F at position 3, S at position 4, T, G, or R at position 5, Y or F at position 6, A, G or T at position 7, M at position 8 and S at position 9;

heavy Chain CDR2 comprises consensus sequence at least 90% identical to sequence: T or A at position 1, I or S at position 2, I or S at position 3, N, S, T or G at position 4, G or R at position 5, G or A at position 6, G, T, or D at position 7, Y, K, H or S at position 8, T or I at position 9, Y or F at position 10, Y at position 11, P or S at position 12 and D at position 13, S or T at position 14, V or L at position 15 and KG for positions 16-17;

heavy chain CDR3 comprises consensus sequence at least 90% identical to sequence: G, L, or N at position 2, G, T, or Y at position 3, G or T at position 4, Y at position 7, Y, A, or G at position 10, M, D or F at position 11, D or E at position 12 and AY at position 14-15;

light chain CDR1 comprises consensus sequence \_ at least 90% identical to sequence: K or R at position 1, A or S at position 2, S or R at position 3, S, Y, I or V at position 8, T or S at position 10, G, S, D, or Q at position 12, V, Y, K or N at position 13, N, S, or T at position 14, Y or F at position 15, and I, L or M at position 16;

light Chain CDR2 comprises consensus sequence at least 90% identical to sequence: A, T or V at position 2, S at position 3, N, T, or K at position 4, L or R at position 5, E, A, F or D at position 6, and S at position 7; and

light chain CDR3 comprises consensus sequence at least 90% identical to sequence: Q, F or W at position 1, H or Q at position 2, R, S, T, Y or N at position 4, E, L, S or H at position 5, L, S, V, D or Y at position 6, P or S at position 7, and T at position 9.

21. The antibody, or fragment thereof of claim 17, which is MNC2, having  
heavy chain CDR1 comprises consensus sequence FTFSGYAMS;  
heavy Chain CDR2 comprises consensus sequence TISSGGTYIYPDSVKG;  
heavy chain CDR3 comprises consensus sequence -LGGDNYEYFDV--;  
light chain CDR1 comprises consensus sequence RASKS--VSTSGYSYMH;  
light Chain CDR2 comprises consensus sequence LASNLES; and  
light chain CDR3 comprises consensus sequence QHSRELPFT.
22. The antibody, or fragment thereof of claim 17, which is MNE6, having  
heavy chain CDR1 comprises consensus sequence FTFSRYGMS;  
heavy Chain CDR2 comprises consensus sequence TISGGGTYYIYPDSVKG;  
heavy chain CDR3 comprises consensus sequence DNYGRNYDYGMDY--;  
light chain CDR1 comprises consensus sequence -----SATSSVSYIH;  
light Chain CDR2 comprises consensus sequence STSNLAS; and  
light chain CDR3 comprises consensus sequence QQRSSSPFT.

23. The antibody, or fragment thereof of claim 17, which is B2, having heavy chain CDR1 comprises consensus sequence FAFSTFAMS; heavy Chain CDR2 comprises consensus sequence AISNGGGYTYYPDTLKG; heavy chain CDR3 comprises consensus sequence ----RYYDLYFDL--; light chain CDR1 comprises consensus sequence RSSQNIV-HSNGNTYLE; light Chain CDR2 comprises consensus sequence KVSNRFS; and light chain CDR3 comprises consensus sequence FQDSHVPLT.
24. The antibody, or fragment thereof of claim 17, which is B7, having heavy chain CDR1 comprises consensus sequence FTFSRYGMS; heavy Chain CDR2 comprises consensus sequence TISSGGTYIYYPDSVKG; heavy chain CDR3 comprises consensus sequence DNYGSSYDYAMDY--; light chain CDR1 comprises consensus sequence RSSQTIV-HSNGNTYLE; light Chain CDR2 comprises consensus sequence KVSNRFS; and light chain CDR3 comprises consensus sequence FQDSHVPLT.
25. The antibody, or fragment thereof of claim 17, which is B9, having heavy chain CDR1 comprises consensus sequence FTFSRYGMS; heavy Chain CDR2 comprises consensus sequence TISSGGTYIYYPDSVKG; heavy chain CDR3 comprises consensus sequence DNYGSSYDYAMDY--; light chain CDR1 comprises consensus sequence -----SASSSVSYM; light Chain CDR2 comprises consensus sequence TTSNLAS; and light chain CDR3 comprises consensus sequence QQRSSYPF-.
26. The antibody, or fragment thereof of claim 17, which is 8C7F3, having heavy chain CDR1 comprises consensus sequence FTFSTYAMS; heavy Chain CDR2 comprises consensus sequence AISNGGGYTYYPDSLKG; heavy chain CDR3 comprises consensus sequence ----RYYDHYFDY--; light chain CDR1 comprises consensus sequence --RASESVATYGNNFMQ; light Chain CDR2 comprises consensus sequence LASTLDS; and light chain CDR3 comprises consensus sequence QQNNEDPPT.
27. The antibody, or fragment thereof of claim 17, which is H11, having

- heavy chain CDR1 comprises consensus sequence FAFSTFAMS;  
heavy Chain CDR2 comprises consensus sequence AISNGGGYTYYPDTLKG;  
heavy chain CDR3 comprises consensus sequence ----RYYDLYFDL--;  
light chain CDR1 comprises consensus sequence RSSQNIV-HSNGNTYLE;  
light Chain CDR2 comprises consensus sequence KVSNRFS; and  
light chain CDR3 comprises consensus sequence FQDSHVPLT.
28. The antibody, or fragment thereof of claim 17, which is B12, having  
heavy chain CDR1 comprises consensus sequence SYGVH;  
heavy Chain CDR2 comprises consensus sequence VIWPGGSTNYNSTLMSRM;  
heavy chain CDR3 comprises consensus sequence DRTPRVGAWFAY; and  
light chain CDR1 comprises consensus sequence RASESVATYGNNFMQ;  
light Chain CDR2 comprises consensus sequence LASTLDS; and  
light chain CDR3 comprises consensus sequence QQNNEDPPT.
29. The antibody, or fragment thereof of claim 17, which is 20A10, having  
heavy chain CDR1 comprises consensus sequence FTFSTYAMS;  
heavy Chain CDR2 comprises consensus sequence -SIGRAGSTYYSDSVKG;  
heavy chain CDR3 comprises consensus sequence ---GPIYNDYDEFAY;  
light chain CDR1 comprises consensus sequence KSSQSVLYSSNQKNYLA;  
light Chain CDR2 comprises consensus sequence WASTRES; and  
light chain CDR3 comprises consensus sequence HQYLSSLT.
30. The antibody, or fragment thereof of claim 17, which is 3C2B1, having  
heavy chain CDR1 comprises consensus sequence ITFSTYTMS;  
heavy Chain CDR2 comprises consensus sequence TISTGGDKTYYSDSVKG;  
heavy chain CDR3 comprises consensus sequence -GTTAMYYYYAMDY;  
light chain CDR1 comprises consensus sequence RASKS---ISTSDYNYIH ;  
light Chain CDR2 comprises consensus sequence LASNLES; and  
light chain CDR3 comprises consensus sequence QHSRELPLT.
31. An antibody, or fragment thereof, for the diagnosis, treatment or prevention of cancers that requires presence of antibody binding conformationally inducing peptide ASRYNLT (SEQ ID NO:1745) of PSMGFR (SEQ ID NO:2).



32. The antibody, or fragment thereof of claim 31, which is 25E6, having  
heavy chain CDR1 comprises consensus sequence FTFSSYGMS;  
heavy Chain CDR2 comprises consensus sequence TISNGGRHTFYPPDSVKG;  
heavy chain CDR3 comprises consensus sequence QTGTEGWFAAY;  
light chain CDR1 comprises consensus sequence KSSQSLLDSDGKTYLN;  
light Chain CDR2 comprises consensus sequence LVSKLDS \_; and  
light chain CDR3 comprises consensus sequence WQGTHFPQT.
33. An antibody, or fragment thereof, for the diagnosis, treatment or prevention of cancers that requires presence of antibody binding conformationally inducing peptide SVSDV (SEQ ID NO:1761) of PSMGFR (SEQ ID NO:2).
34. The antibody of claim 33, which is 5C6F3, having  
heavy chain CDR1 comprises consensus sequence FTFSTYAMS ;  
heavy Chain CDR2 comprises consensus sequence AISNGGGYTYYPDSLKG;  
heavy chain CDR3 comprises consensus sequence RYYDHYFDY;  
light chain CDR1 comprises consensus sequence RSSQTIVHSNGNTYLE;  
light Chain CDR2 comprises consensus sequence KVSNRFS; and  
light chain CDR3 comprises consensus sequence FQDSHVPLT.
35. The antibody or fragment thereof according to claims 1-34, which is murine, camelid, human or humanized.
36. The antibody or fragment thereof according to claims 1-34, wherein the antibody fragment is scFv or scFv-Fc, which variable regions thereof may be murine, camelid, human or humanized.
37. A chimeric antigen receptor (CAR) comprising the antibody fragment of claim 36.
38. A chimeric antigen receptor (CAR) comprising the antibody fragment of claim 36, further comprising mutations in the co-stimulatory domain or CD3-zeta signaling domain.
39. The CAR of claim 38, wherein tyrosines are mutated in CD28 or 4-1BB.
40. The CAR of claim 38, wherein the CD3-zeta contains 1XX mutations.
41. An immune cell comprising the CAR of claim 37.
42. The immune cell of claim 41, which is a T cell, NK cell, dendritic cell, or mast cell.

43. A cell composition expressed in a cell comprising a CARs of claim 37, and second entity having a biological recognition unit that has a specificity that is different from that of the CAR.
44. The composition of claim 43, wherein the second entity binds PD-1, PDL-1, or other checkpoint inhibitor.
45. The composition of claim 43, wherein the second entity binds to NME7.
46. The composition of claim 43, wherein the second entity is a cytokine.
47. The composition of claim 46, wherein the cytokine is IL-12.
48. The composition of claim 46, wherein the cytokine is IL-18
49. The composition of claim 43, wherein the second entity is c-Jun
50. An immune cell engineered to express a nucleic encoding a CAR of claim 37 and a nucleic acid encoding a second entity as in any of the claims above wherein the second entity expressed from an inducible promoter.
51. The immune cell of claim 50, wherein the second entity is expressed from an inducible promoter that is activated by elements of an activated immune cell.
52. The immune cell of claim 51, wherein the second entity is expressed from an NFAT inducible promoter.
53. The immune cell of claim 52, wherein the NFAT is NFATc1, NFATc3 or NFATc2.
54. The immune cell of claim 50, wherein the second entity is a cytokine.
55. The immune cell of claim 51, wherein the cytokine is IL-7, IL-15, or IL-18.
56. The immune cell of claim 50, wherein the nucleic acids encoding the second entity are inserted into a Foxp3 promoter or enhancer region.
57. The immune cell of claim 56, wherein the cytokine is IL-18.
58. The immune cell of claim 56, wherein the cytokine is expressed from an NFAT inducible promoter.
59. A BiTE construct comprising the antibody fragment of claim 36.
60. An antibody drug conjugate (ADC) comprising the antibody or antibody fragment of claim 35.
61. An antibody or fragment thereof that specifically binds to PSMGFR (SEQ ID NO:2) and N-10 (SEQ ID NO:3); and
  - (i) does not bind to full-length MUC1;
  - (ii) does not bind to C-10 (SEQ ID NO:825);

- (iii) competitively inhibits binding of NME1 or NME7<sub>AB</sub> to MUC1\* extra cellular domain or a PSMGFR peptide;
- (iv) recognizes a MUC1\* generated by cleavage by a cleavage enzyme;
- (v) recognizes a conformational epitope and not a linear epitope; or
- (vi) is cancer selective by immunohistochemistry on tissues.

62. The antibody or fragment thereof of claim 61, wherein four of the criteria (i) – (vi) are satisfied.

63. The antibody or fragment thereof of claim 61, wherein five of the criteria (i) – (vi) are satisfied.

64. The antibody or fragment thereof of claim 61, wherein six of the criteria (i) – (vi) are satisfied.

65. The antibody or fragment thereof of claim 61, wherein at least criteria (vi) is satisfied.

66. The antibody or fragment thereof of claim 61, wherein cleavage enzyme is MMP-9.

67. The antibody or fragment thereof of claims 1-66, wherein the cancer is breast cancer, pancreatic cancer, ovarian cancer, lung cancer, colon cancer, gastric cancer or esophageal cancer.



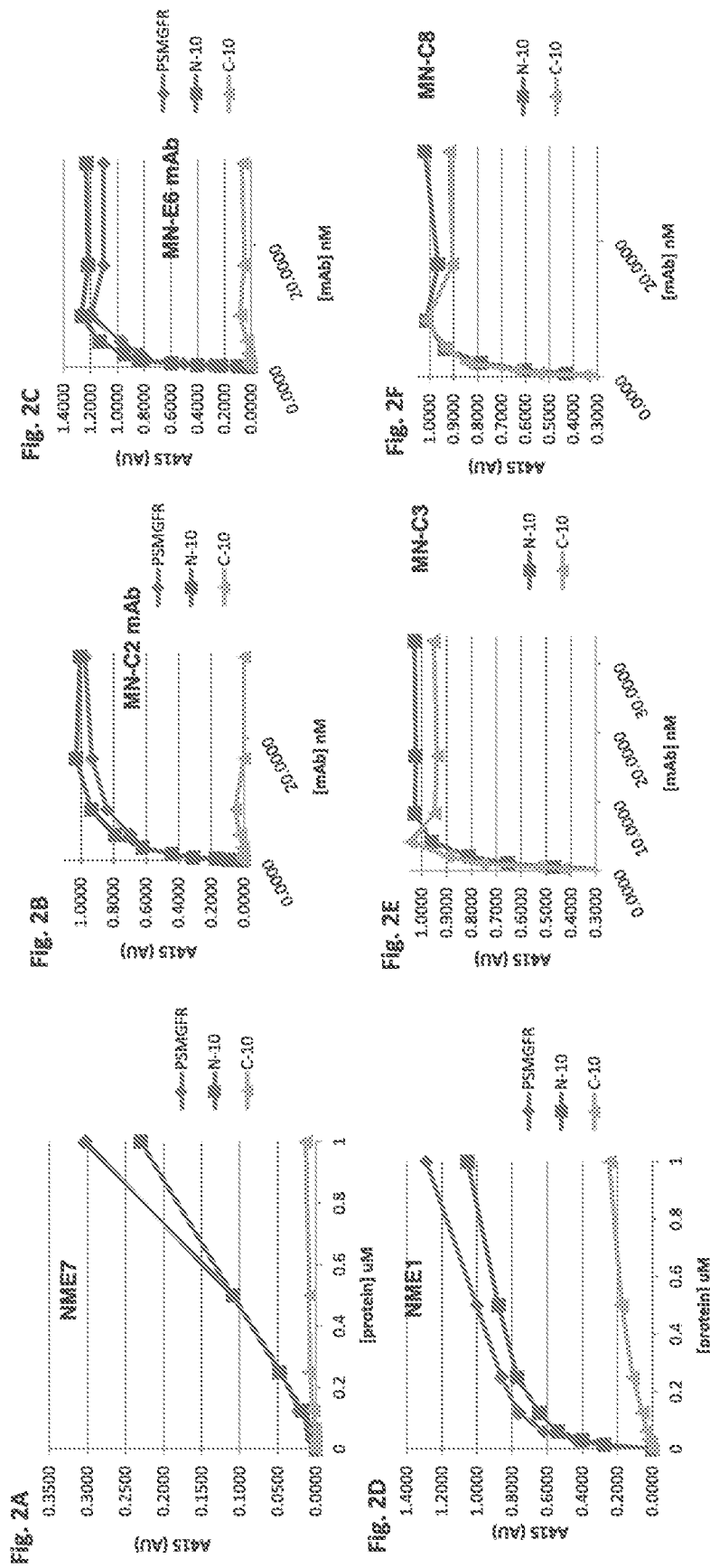
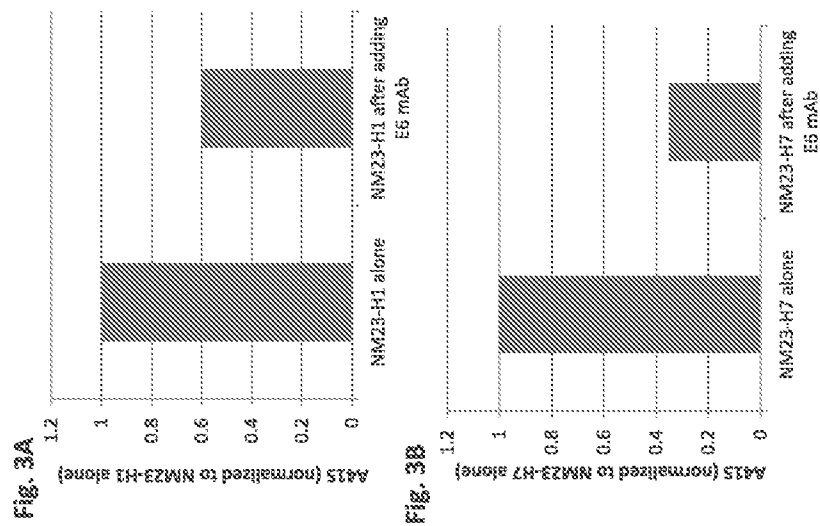
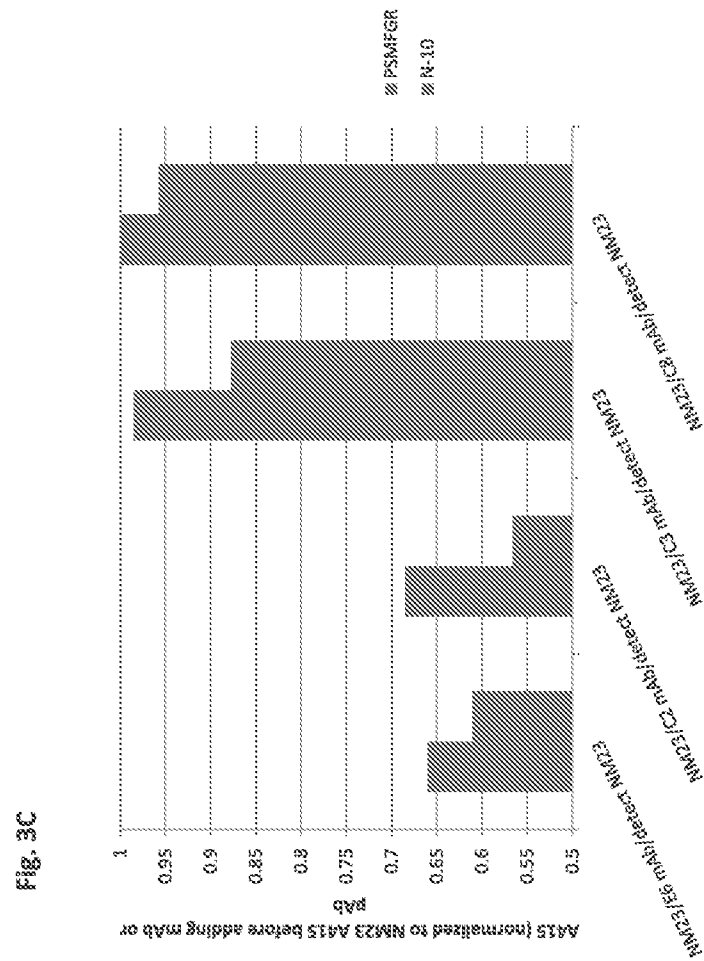


Figure 2A-2I

Fig. 2G PSMGFR GTINVDVETQFNQYKTEAASRYNLTISDVSVSDVPPFSAQSGA

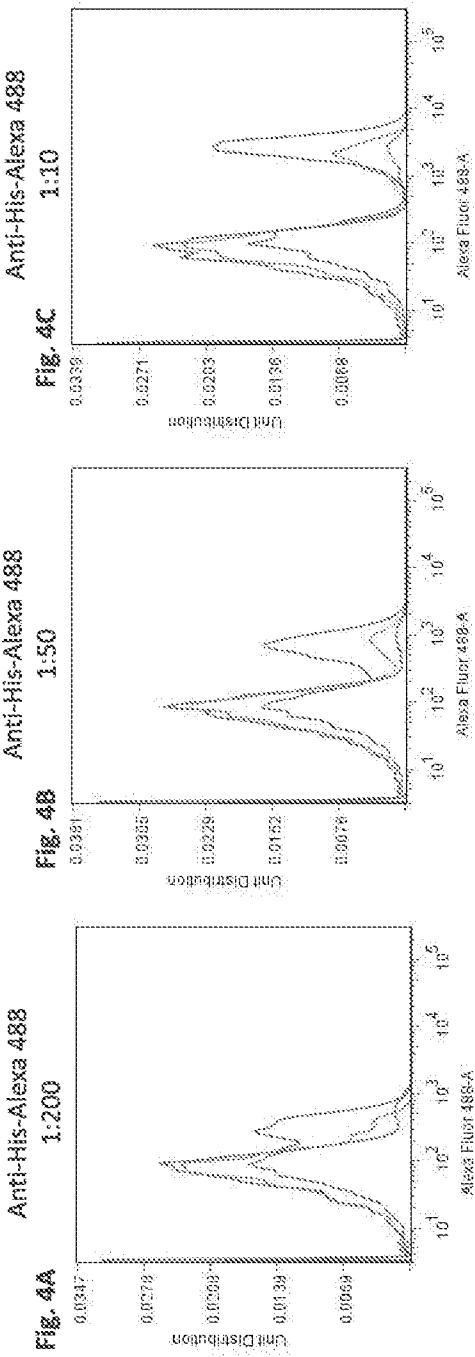
Fig. 2H PSMGFR N-10 QFNQYKTEAASRYNLTISDVSVSDVPPFSAQSGA

Fig. 2I PSMGFR C-10 GTINVDVETQFNQYKTEAASRYNLTISDVSVSDV

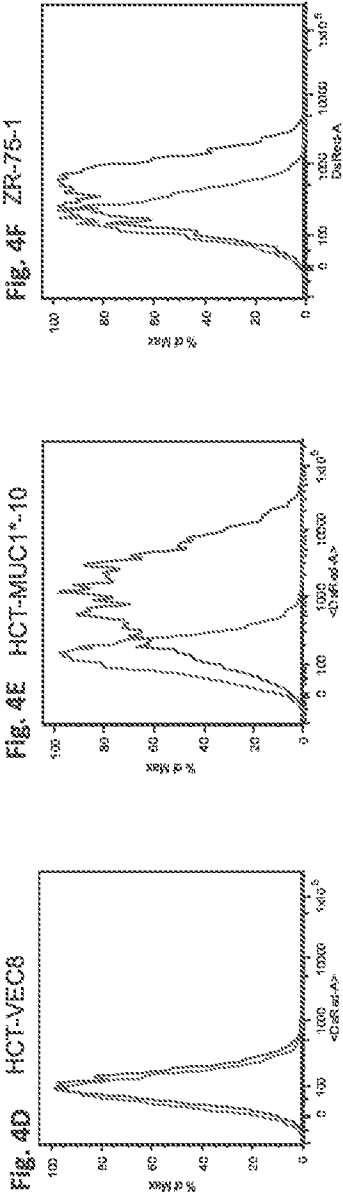


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Humanized MN-C2 scFv binds to ZR-75-1, aka 1500, MUC1\* positive breast cancer cells



Red No scFv  
Green human-MN-C2 scFv 1:10  
of 1.5mg/ml  
Blue scFv 1:2 of 1.5 mg/ml



Red Anti-mouse-PE alone  
Blue MN-E6 Antibody

Figure 4A-4F

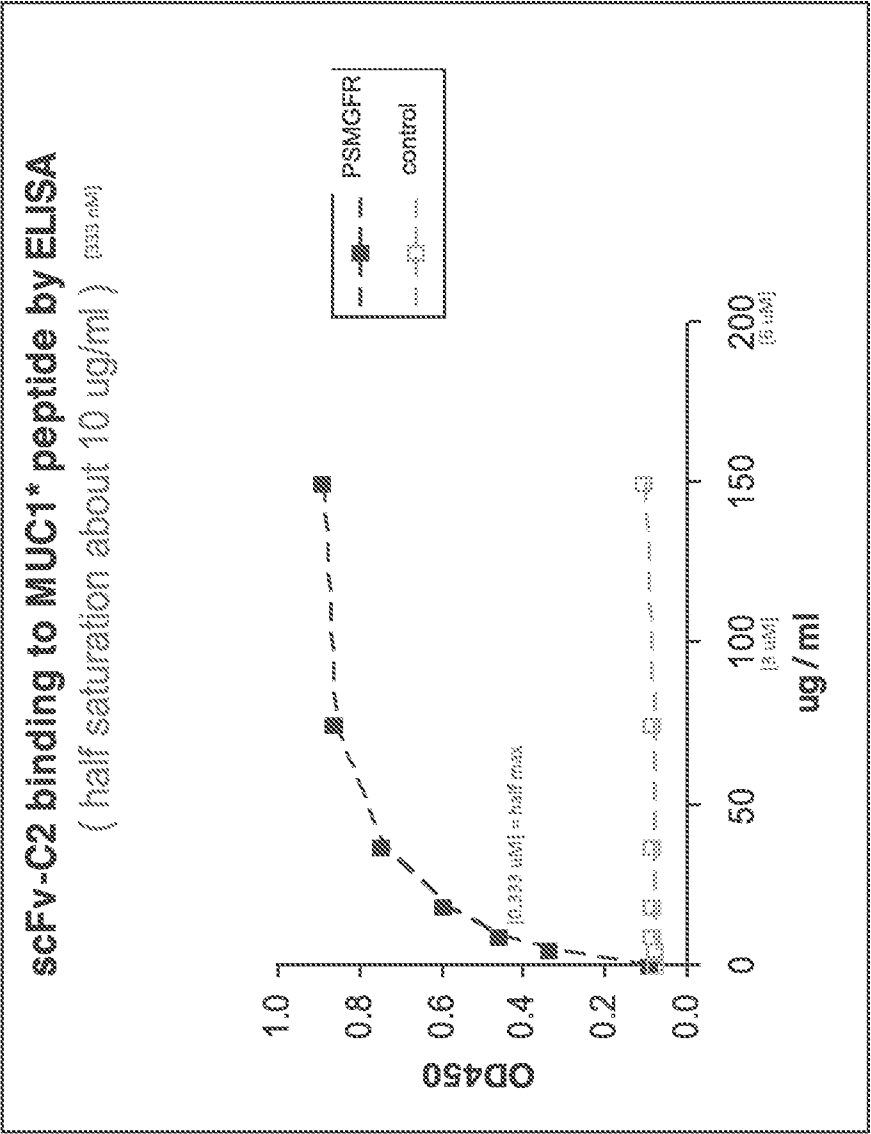


Figure 5



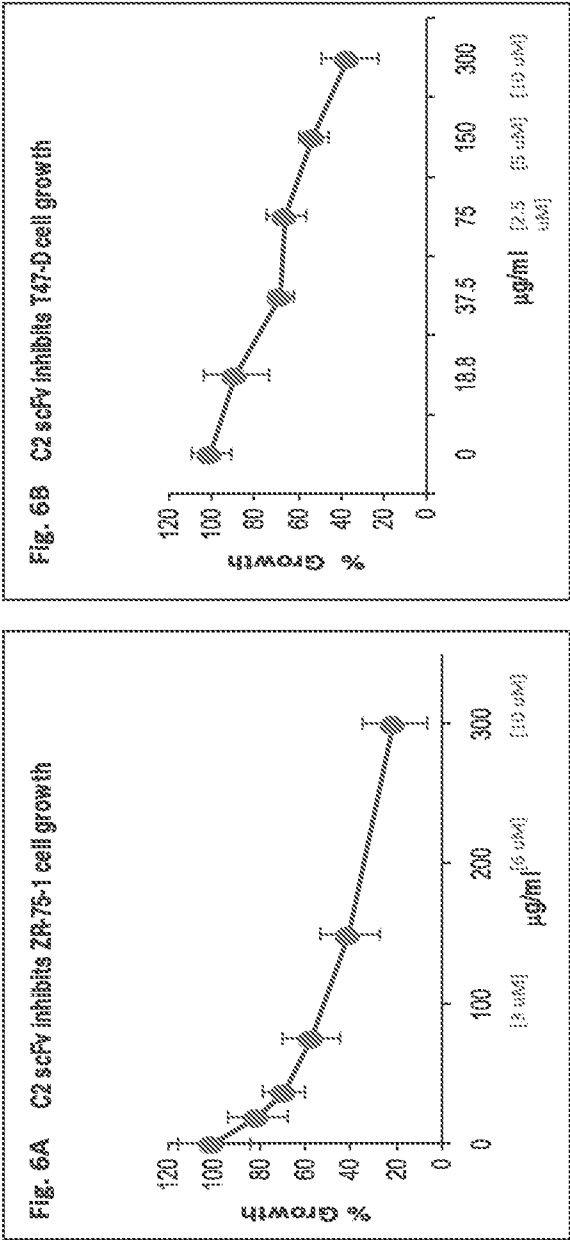


Figure 6A-6B

7/258

Fig. 7A

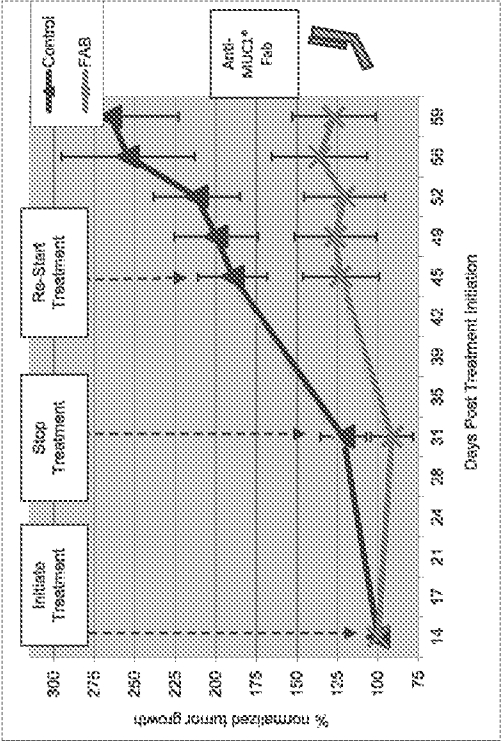


Fig. 7B

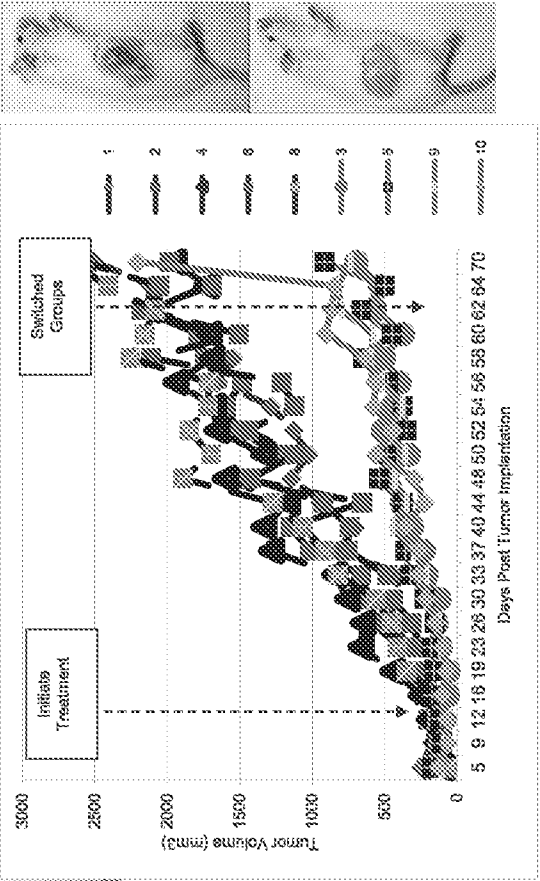


Figure 7A-7B

ELISA – hu MN-E6 scFv-Fc binding to MUC1\* peptides PSMGFR, PSMGFR N-10 and PSMGFR C-10

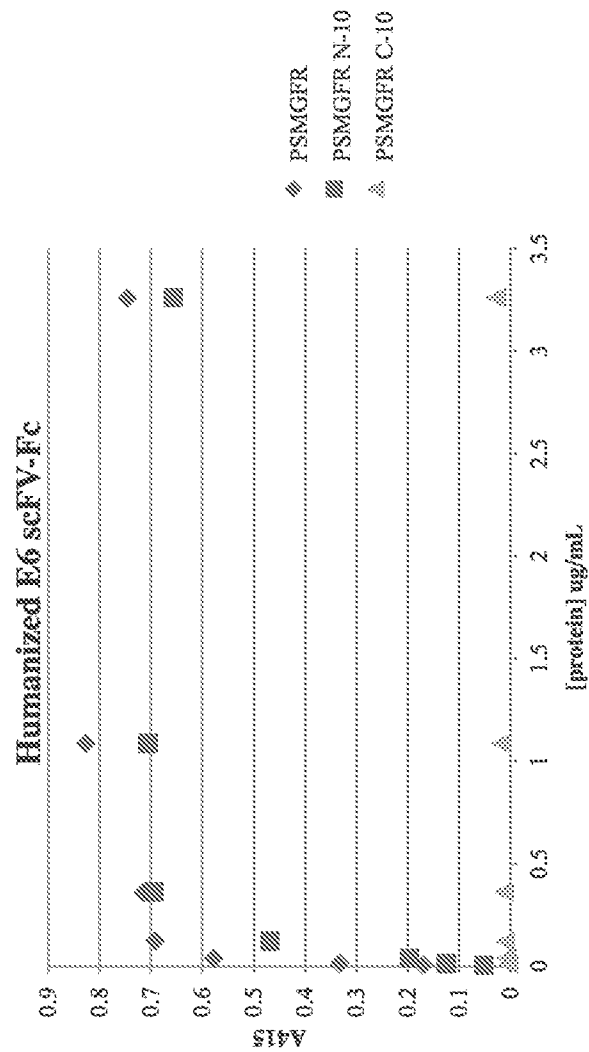


Figure 8

ELISA – mouse monoclonal MN-C3, humanized MN-C3 and humanized MN-C3 scFv-Fc binding to MUC1\* peptides PSMGFR, PSMGFR N-10 and PSMGFR C-10

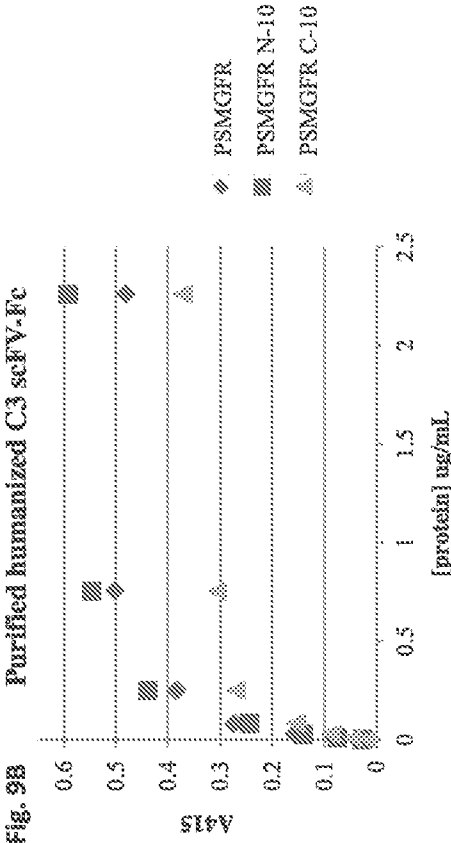
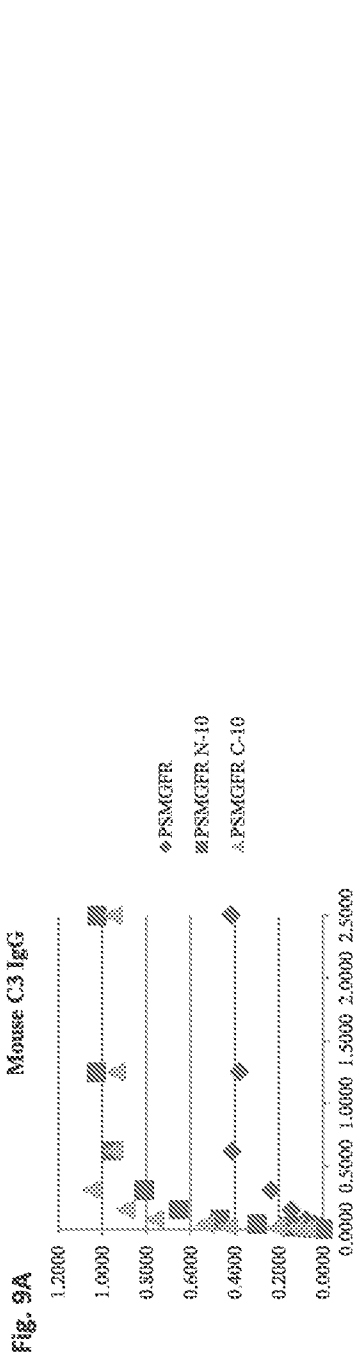
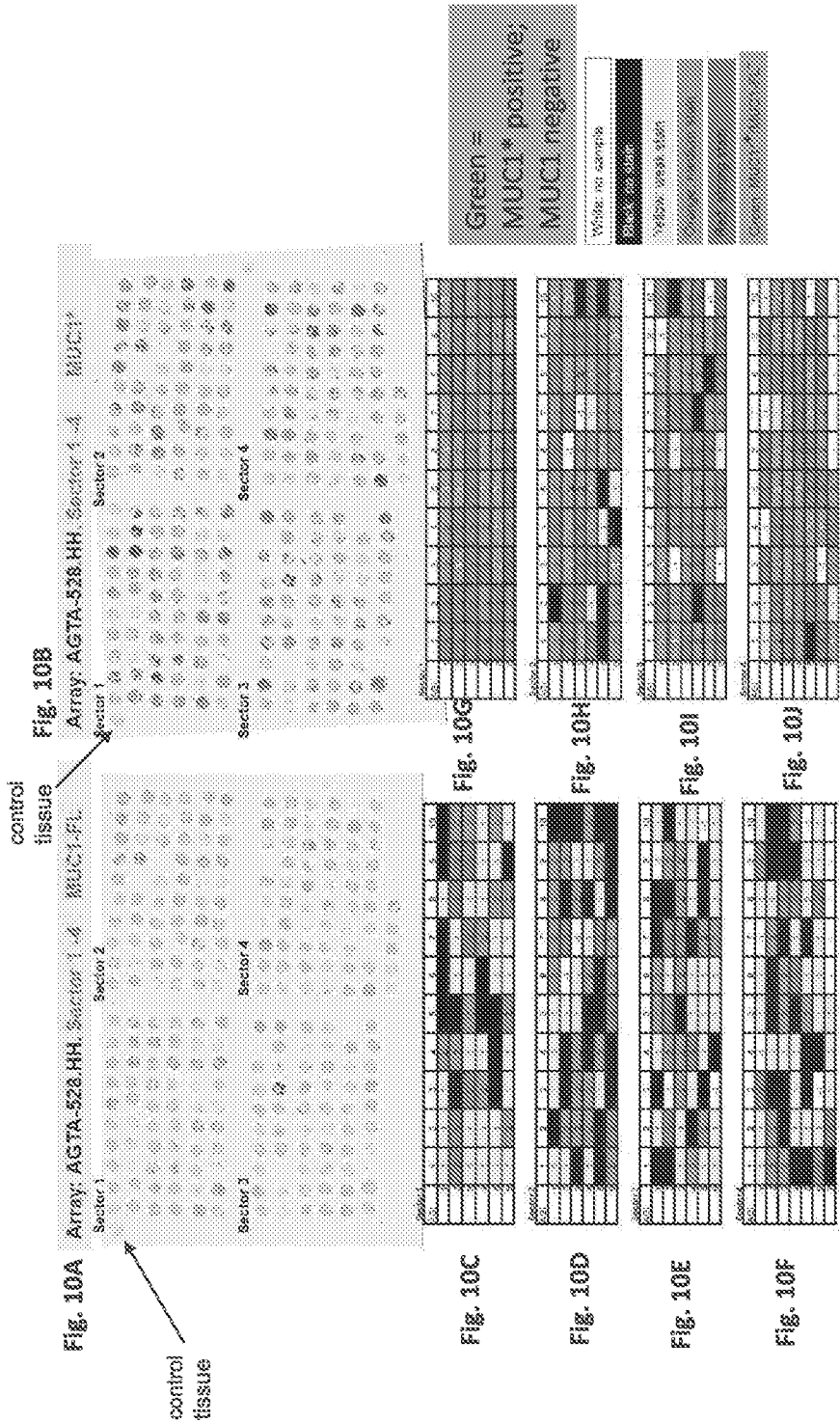
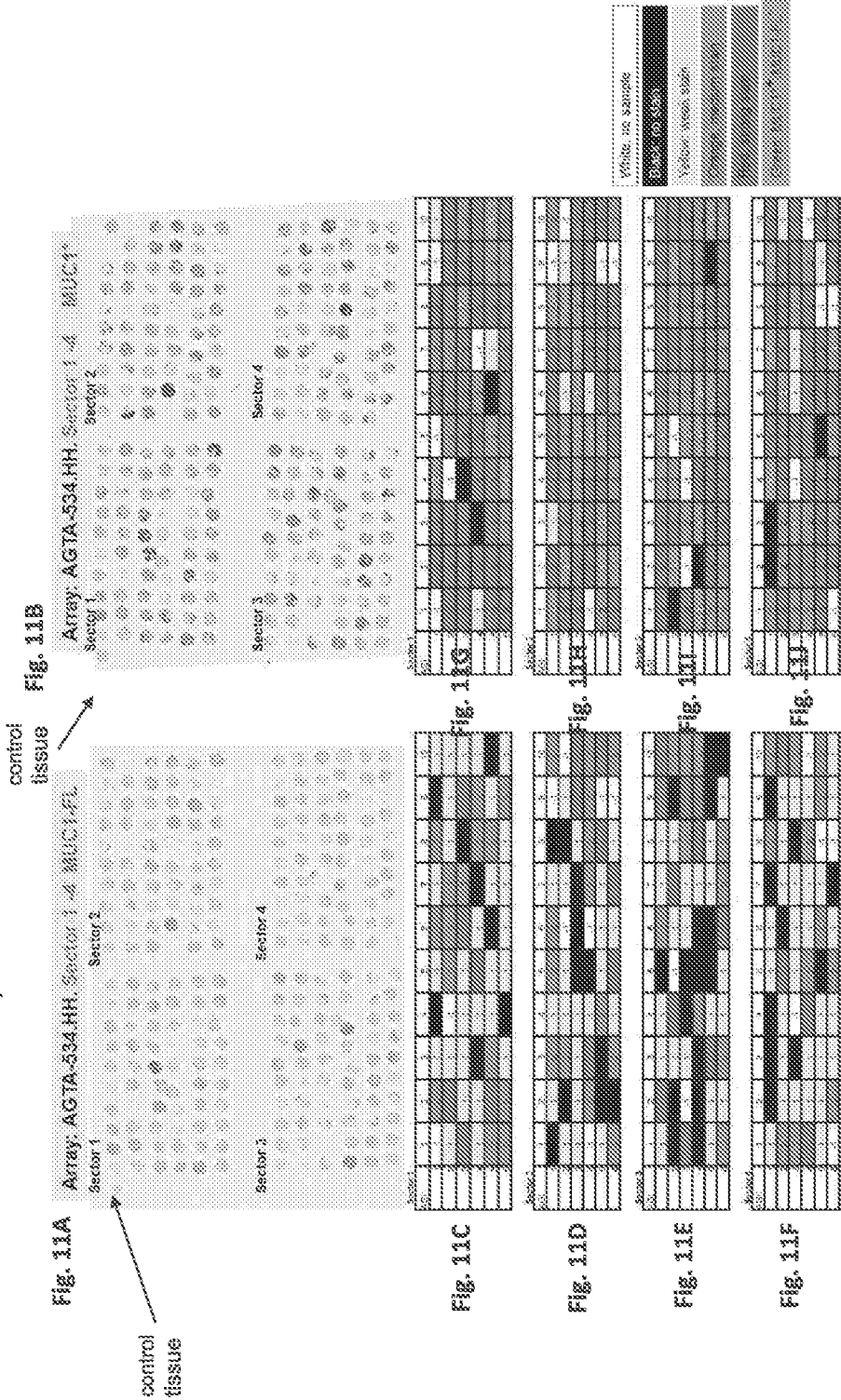


Figure 9A-9B

Tissue arrays comprising specimens from 240 breast cancer patients were stained with an antibody (VU4H5) that recognizes full-length MUC1 (left) or stained with an antibody that recognizes MUC1\* (MN-C2). The data show that most or all (green boxes) of the MUC1 on cancerous tissue is MUC1\* and not MUC1 full-length (MUC1-FL). The data further show that MN-C2 monoclonal antibody binds to cancerous tissue but not the healthy control tissue.



Tissue arrays comprising specimens from 240 breast cancer patients were stained with an antibody (VU4H5) that recognizes full-length MUC1 (left) or stained with an antibody that recognizes MUC1\* (MN-C2). The data show that most or all (green boxes) of the MUC1 on cancerous tissue is MUC1\* and not MUC1 full-length (MUC1-FL). The data further show that MN-C2 monoclonal antibody binds to cancerous tissue but not the healthy control tissue.



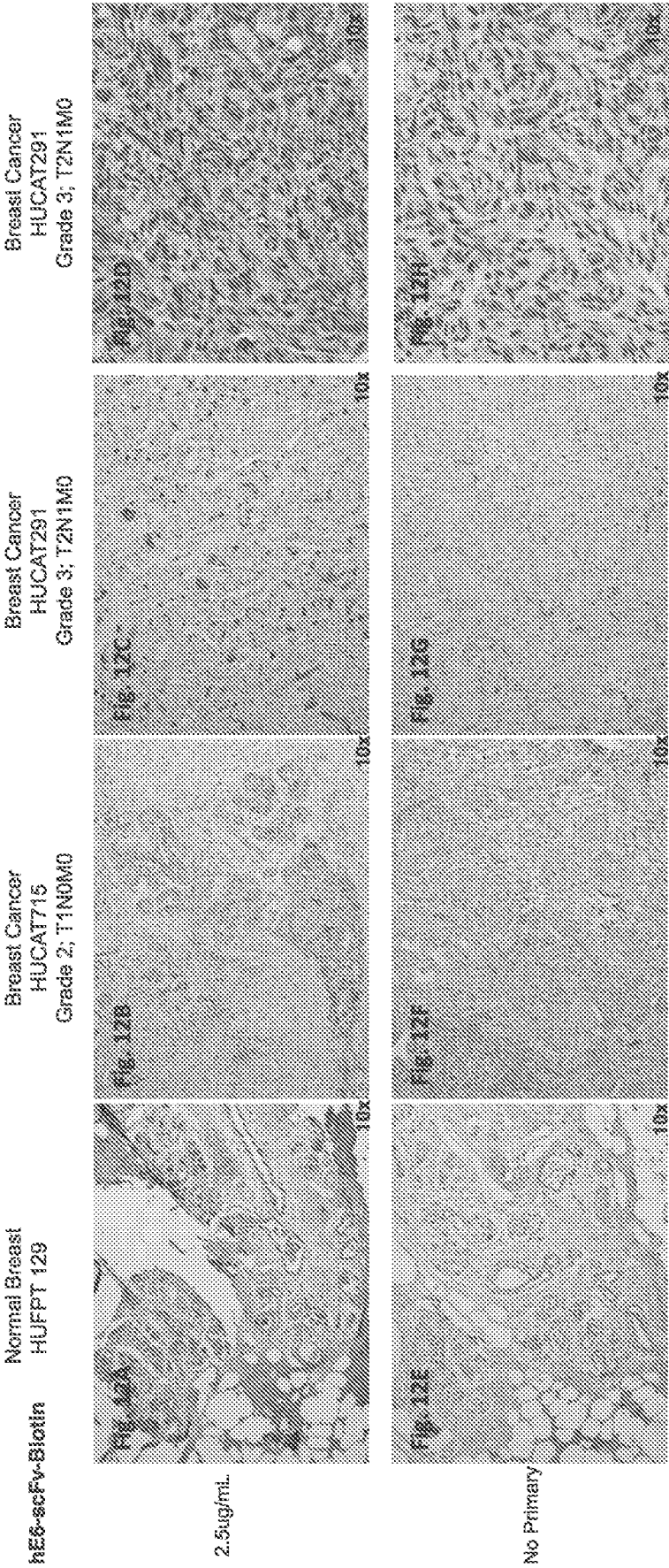


Figure 12A-12H

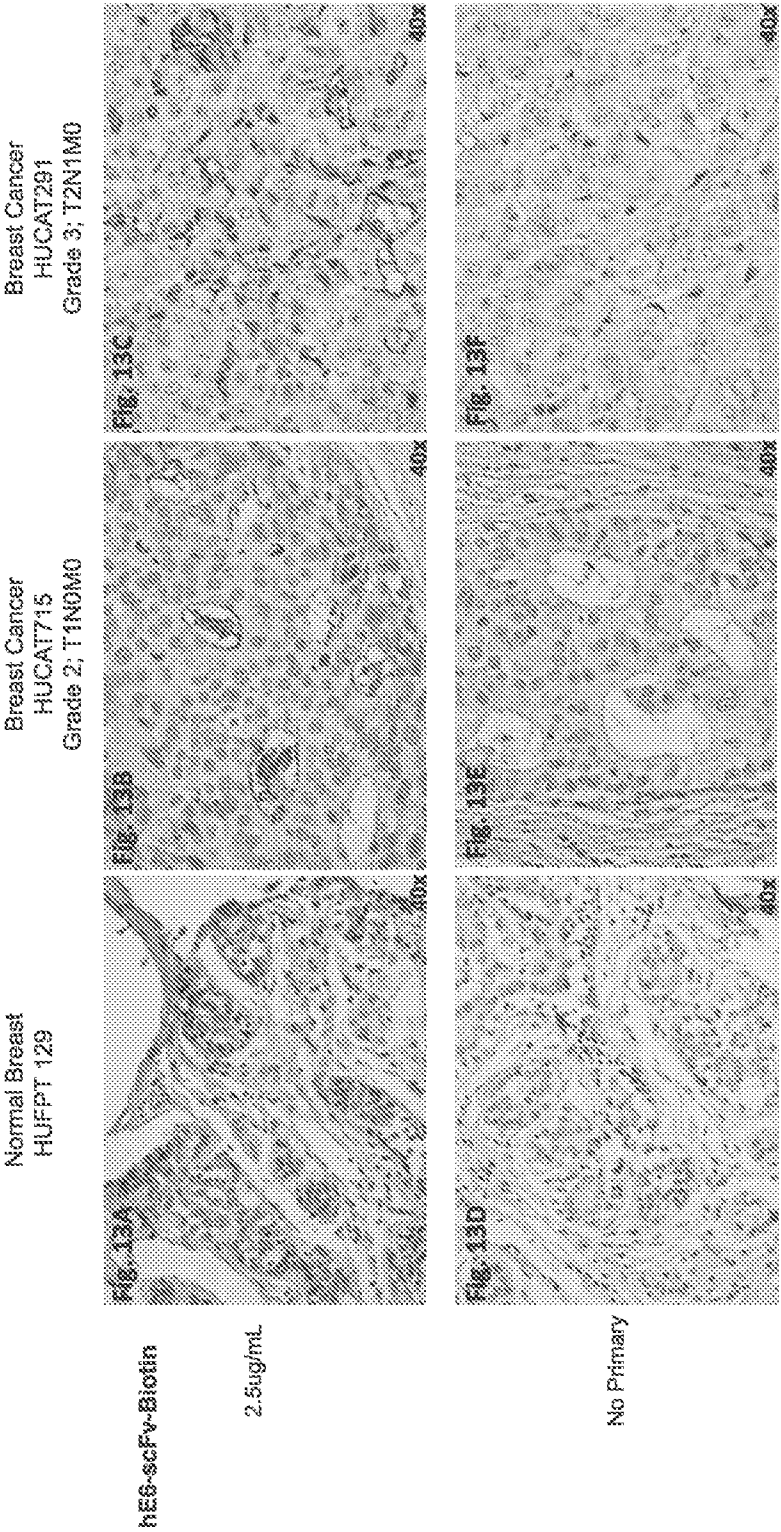
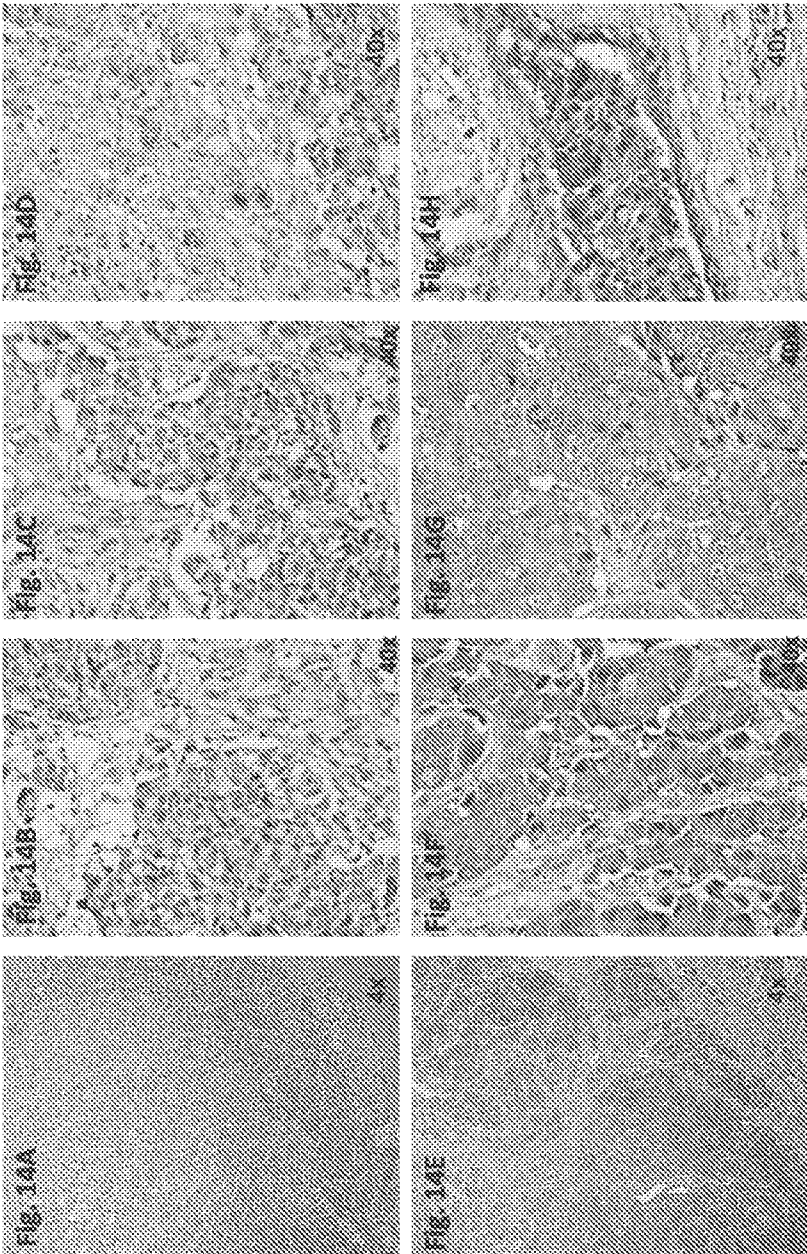


Figure 13A-13F

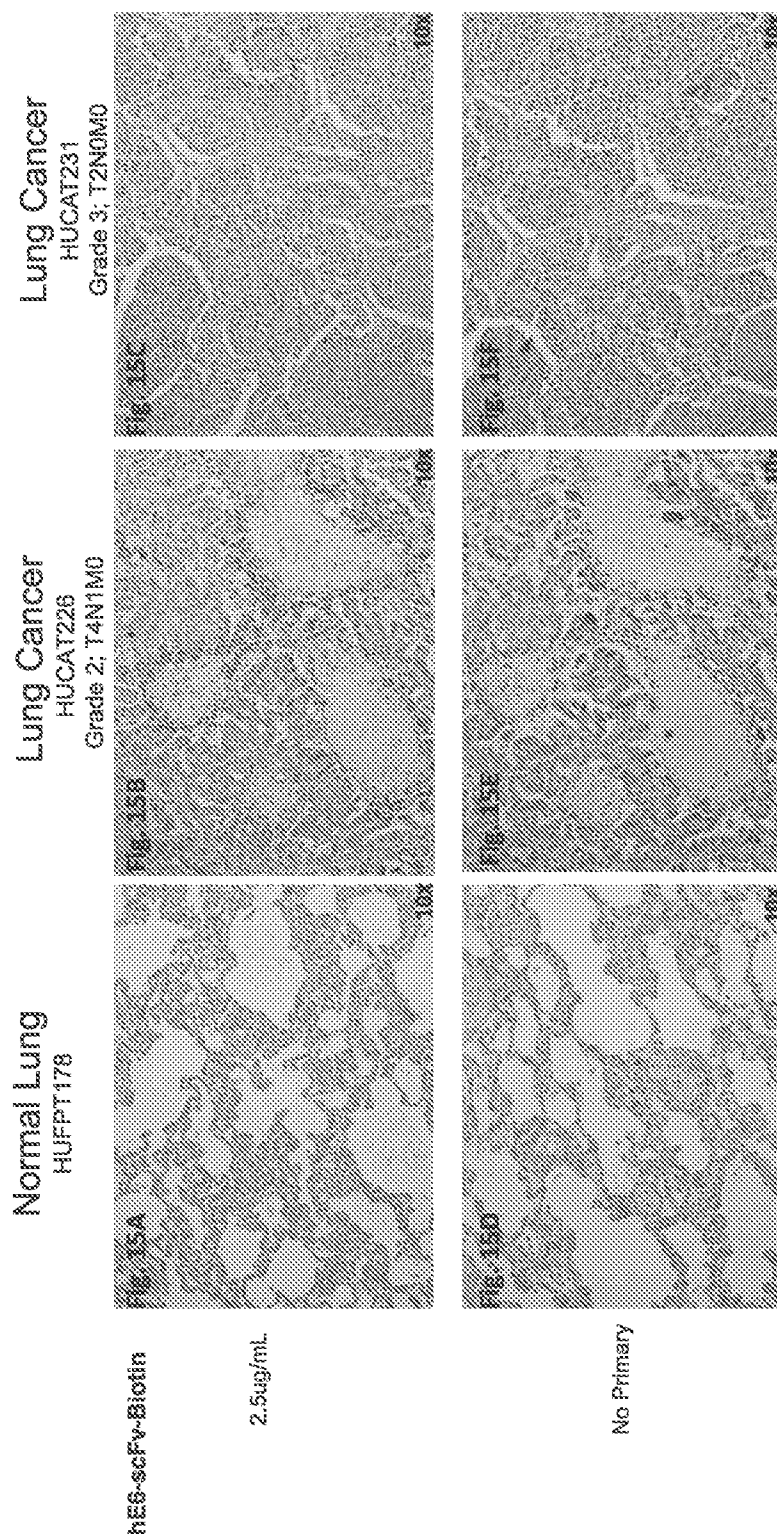




MN-E6  
10 ug/ml  
Breast Cancer  
HUCAT300  
Grade 3

Breast Cancer  
HUCAT291  
Grade 3; T2N1M0

Figure 14A-14H



451-551

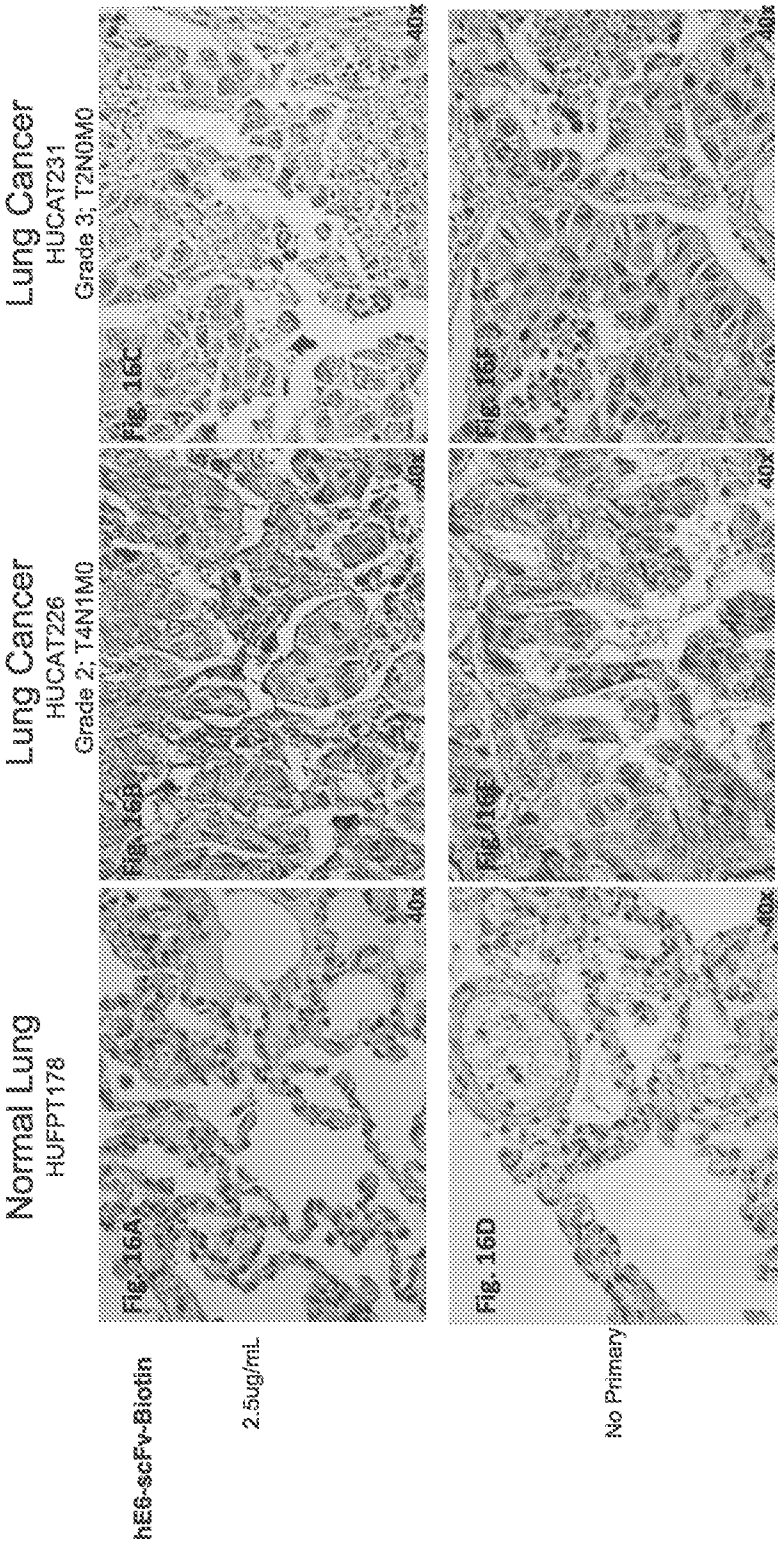


Figure 16A-16F

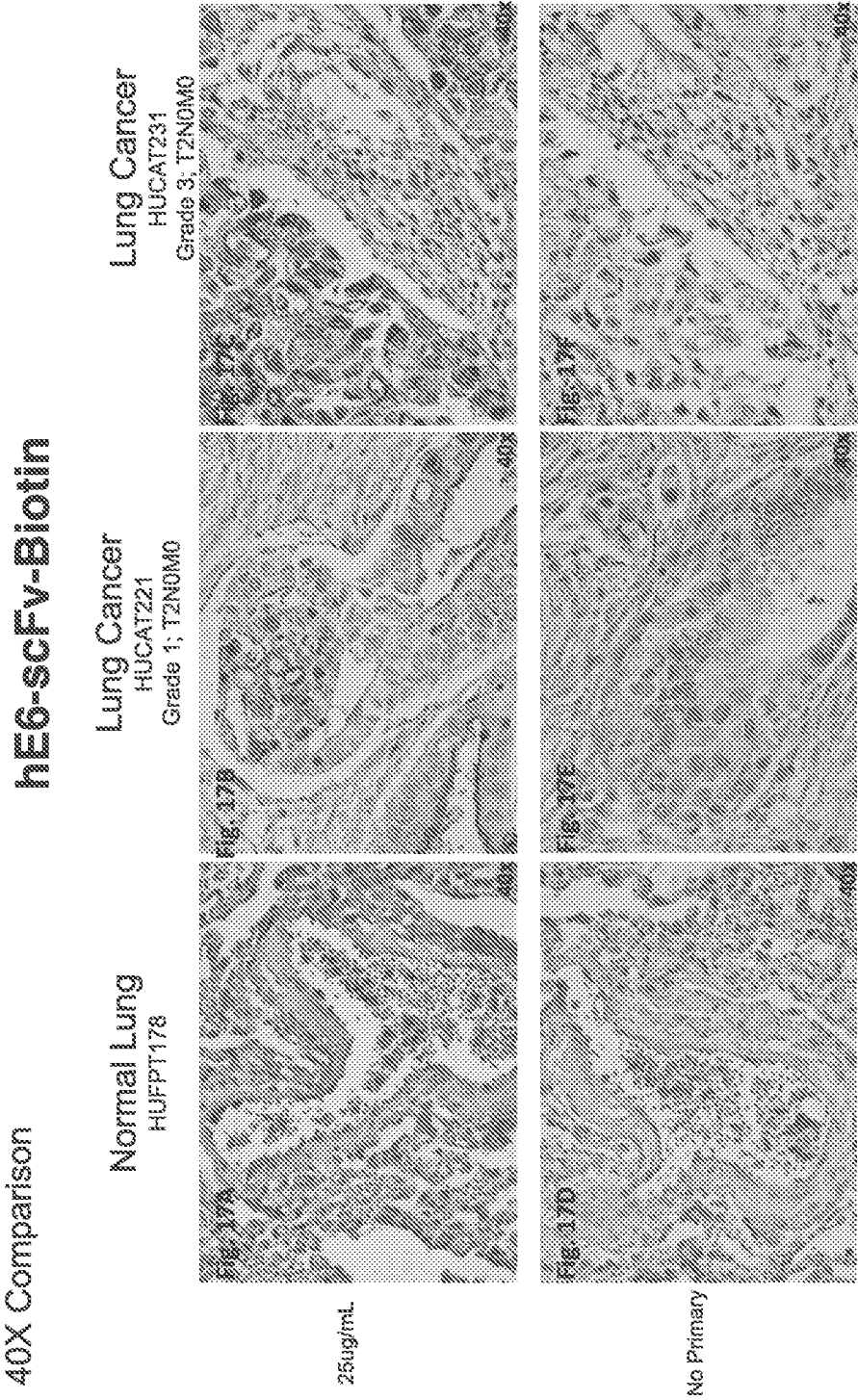


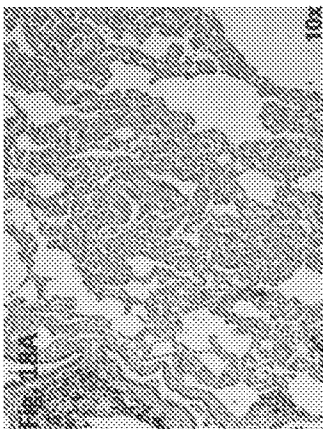
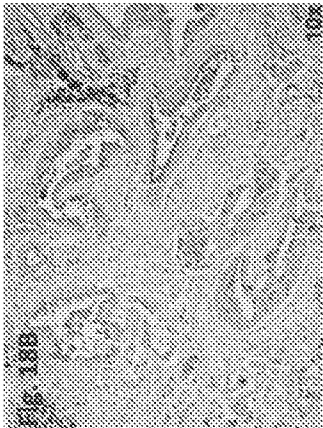
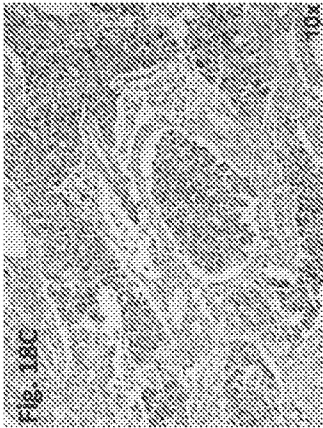
Figure 17A-17F

**hE6-scFv-Biotin**

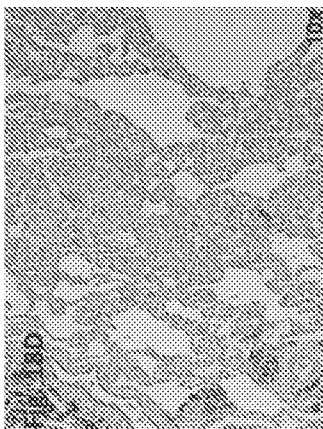
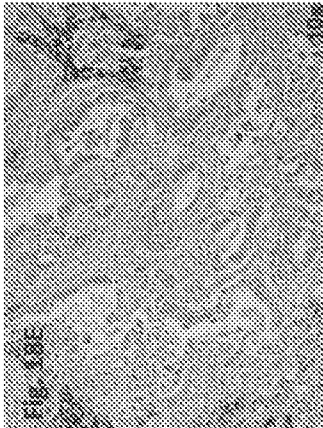
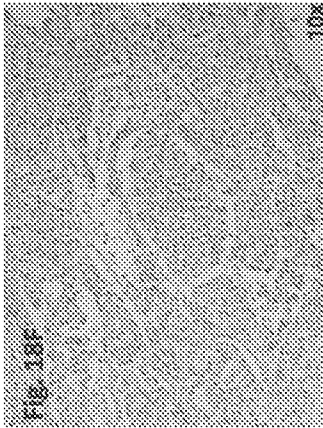
**Lung Cancer**  
BCS040117a E12  
Grade 3; T4N2M0

**Lung Cancer**  
BCS040117a A3  
Grade 1; T2N0M0

**Normal Lung**  
HUFPT178



25ug/mL

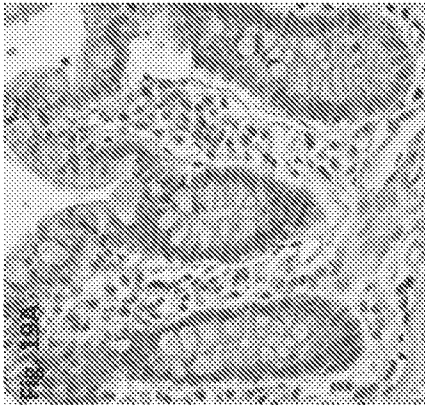


No Primary

Figure 18A-18F

Normal Small Intestine

HUPPT081



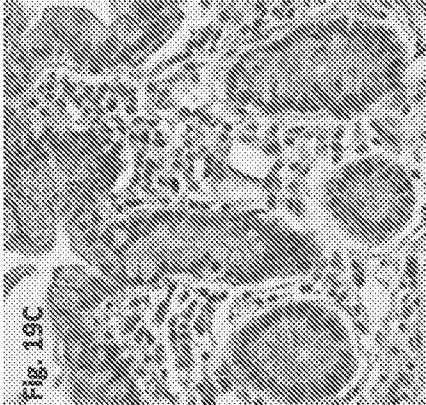
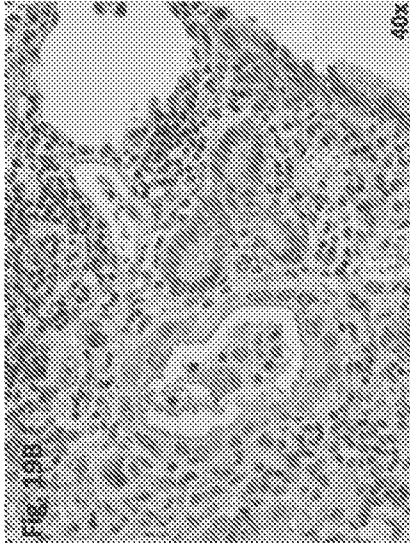
hu-E6-scFv-Fc-  
biotin

5ug/mL

Small Intestine Cancer

HUCAT101m

Grade 3; T3N0M0



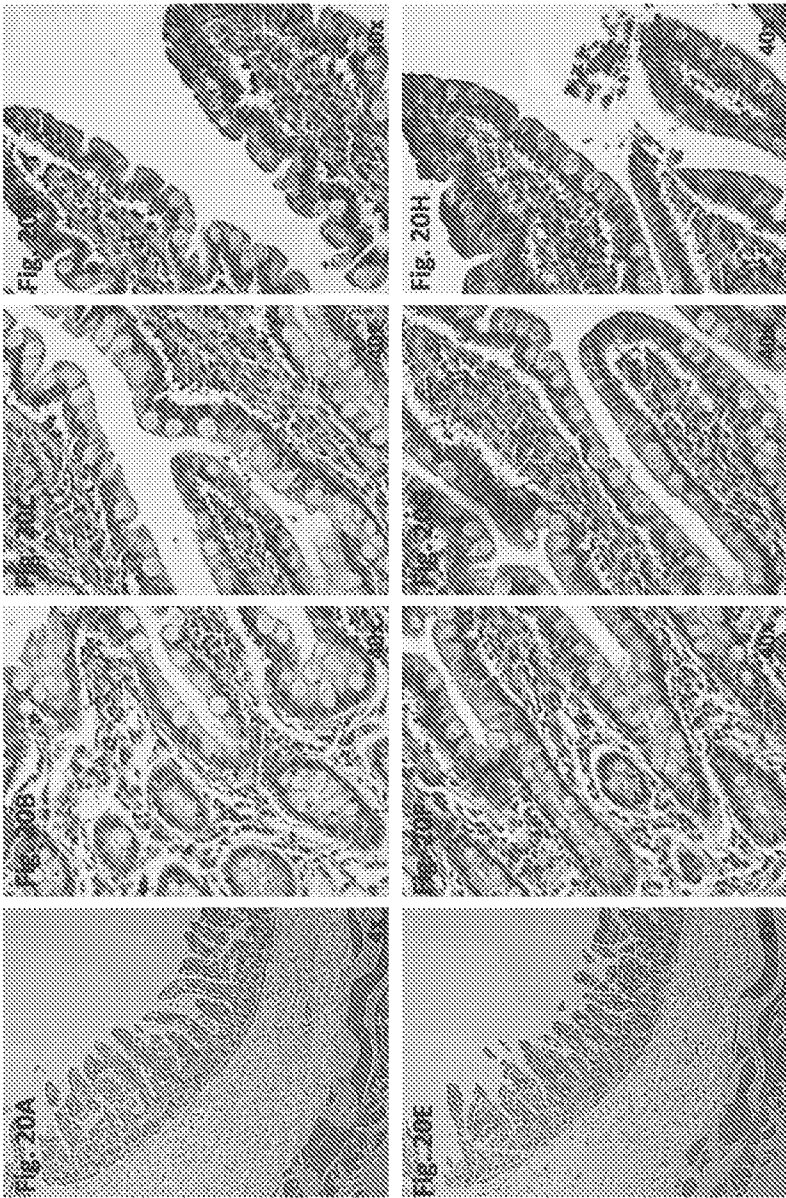
No Primary



Figure 19A-19D



Small Intestine Normal  
HUFPT081

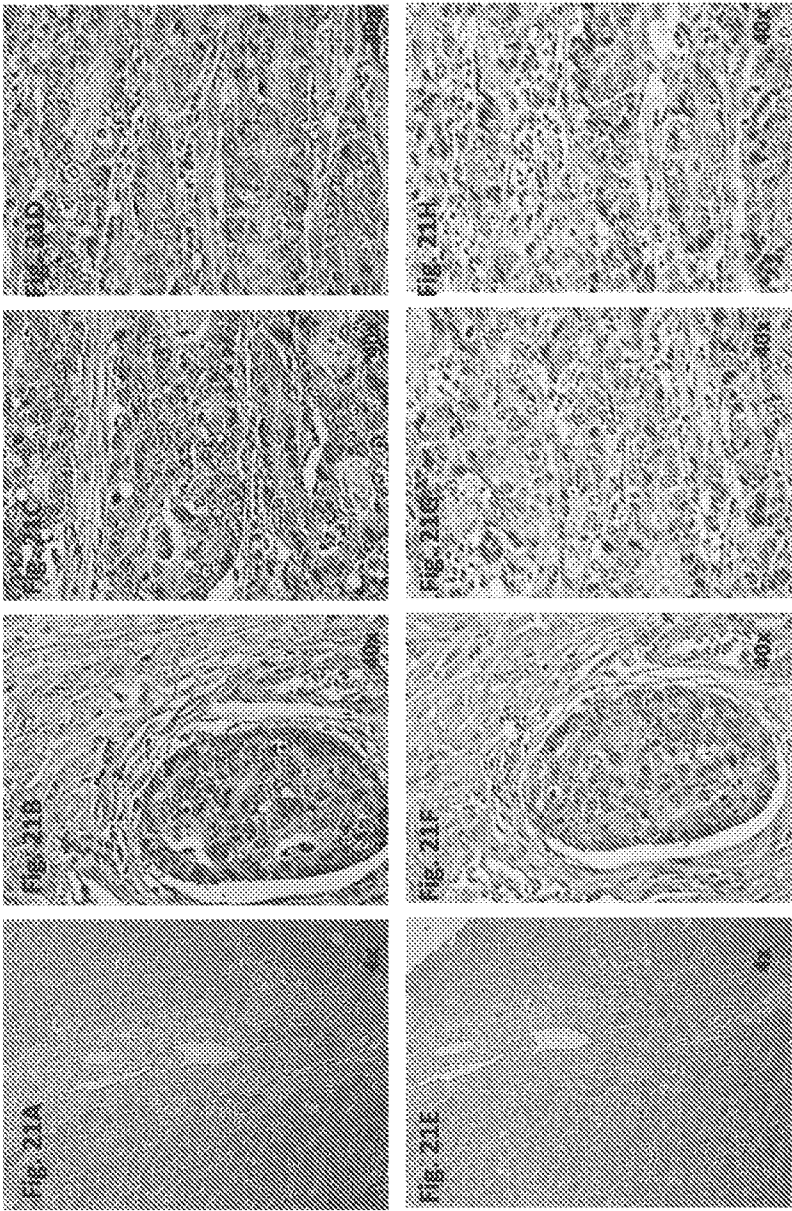


hu-E8  
scFv+Fc  
50 ug/ml  
2' (1:1000)

2' only control  
(GaH-HRP)  
1:1000

Figure 20A-20H

Small Intestine Cancer  
HUCAT096  
Grade 3; T3N0M0



hu-E6  
scFv+Fc

50 ug/ml  
2' (1:1000)

2' only control  
(GaH-HRP)

1:1000

Figure 21A-21H



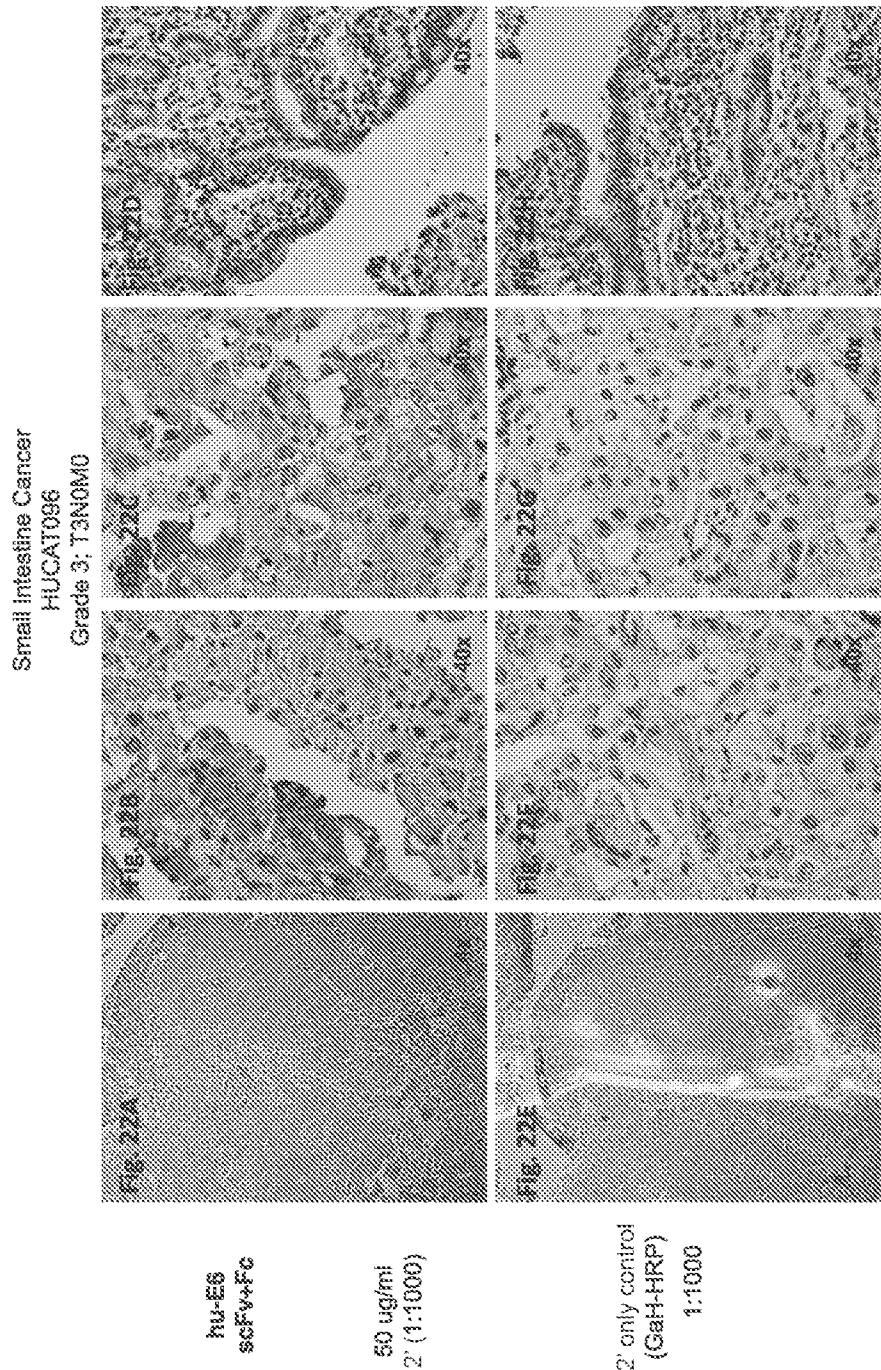
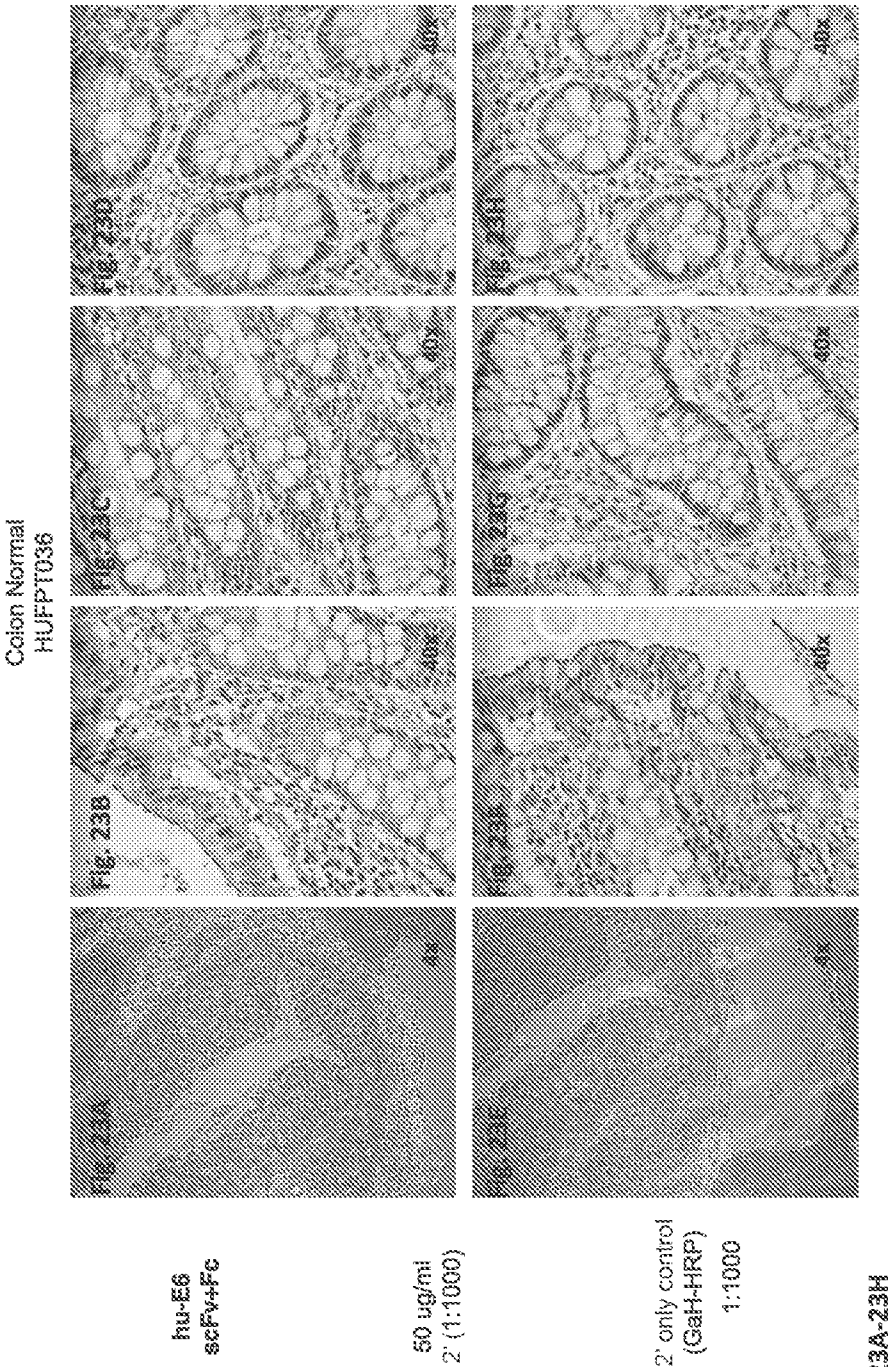
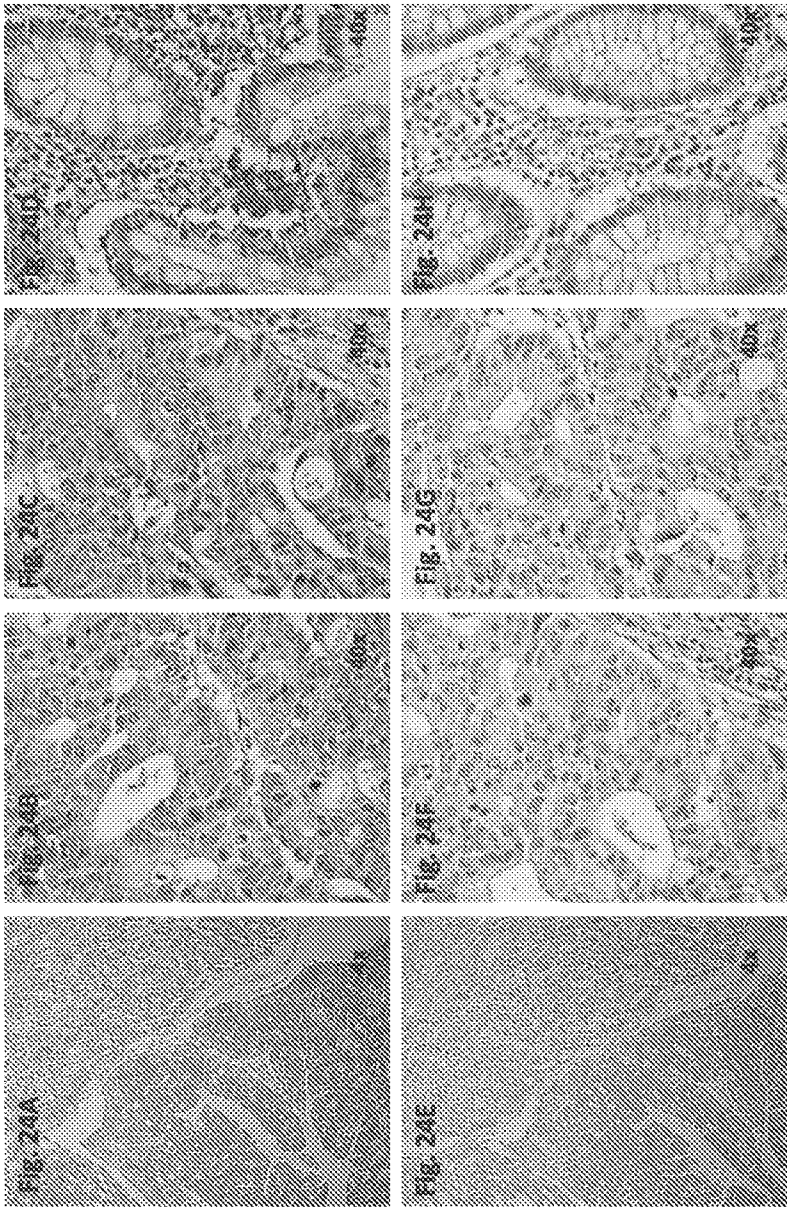


Figure 22A-22H



Colon Cancer  
HUCAT116  
Grade 1; T4N1M1

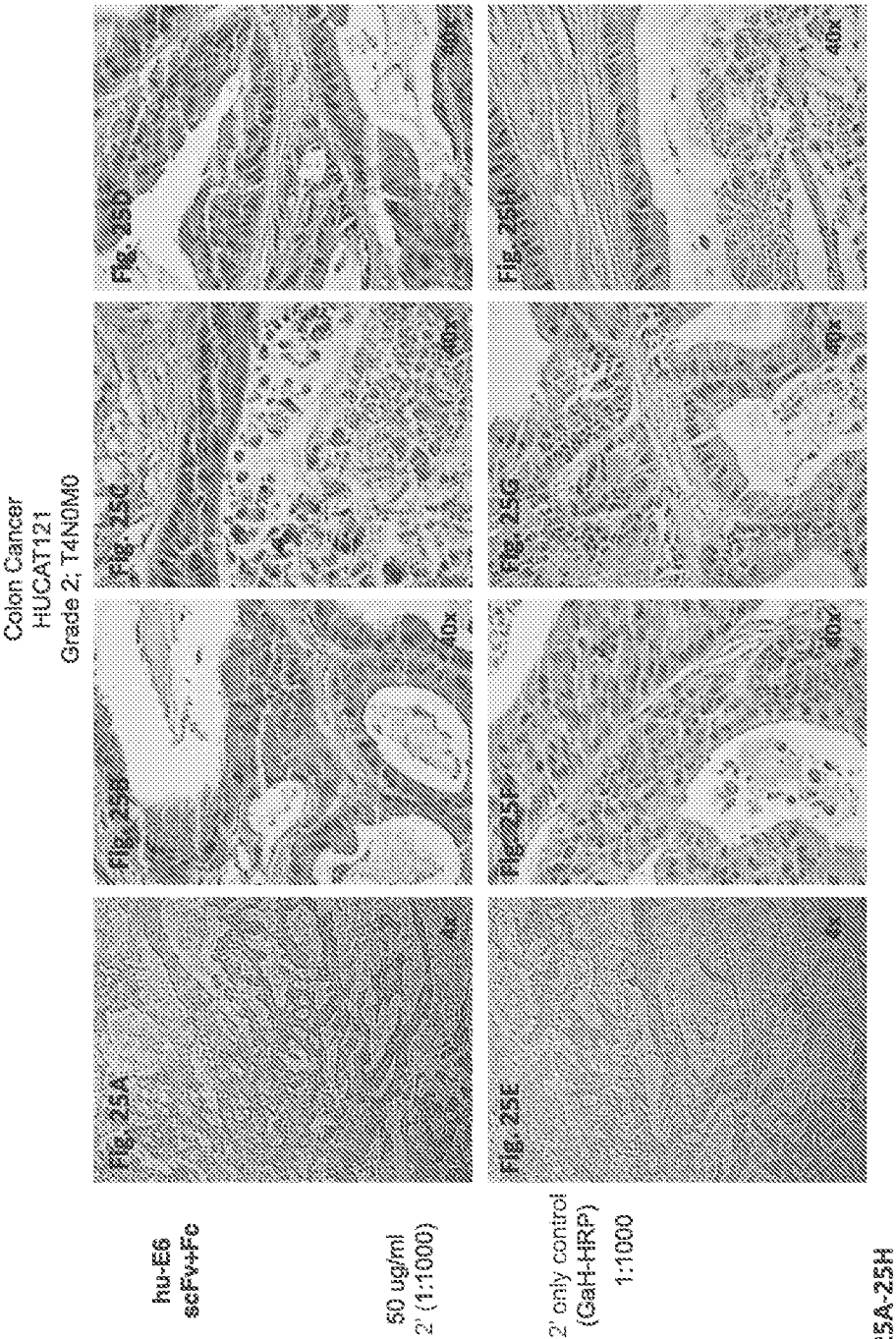


hu-E6  
scFv4Fc

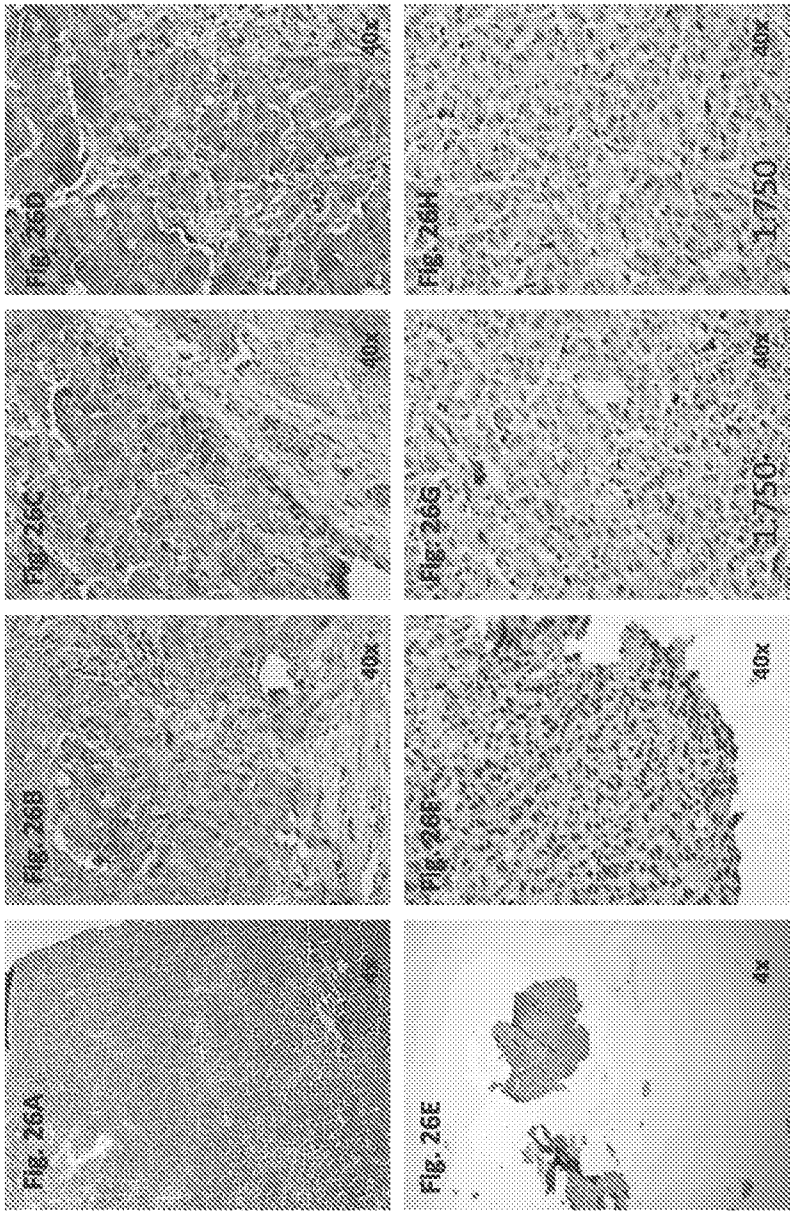
50 ug/ml  
2' (1:1000)

2' only control  
(GaH-HRP)  
1:1000

Figure 24A-24H



Colon Cancer  
HUCAT128  
Grade 3; T3N1M1



hu-E6  
scFv+Fc

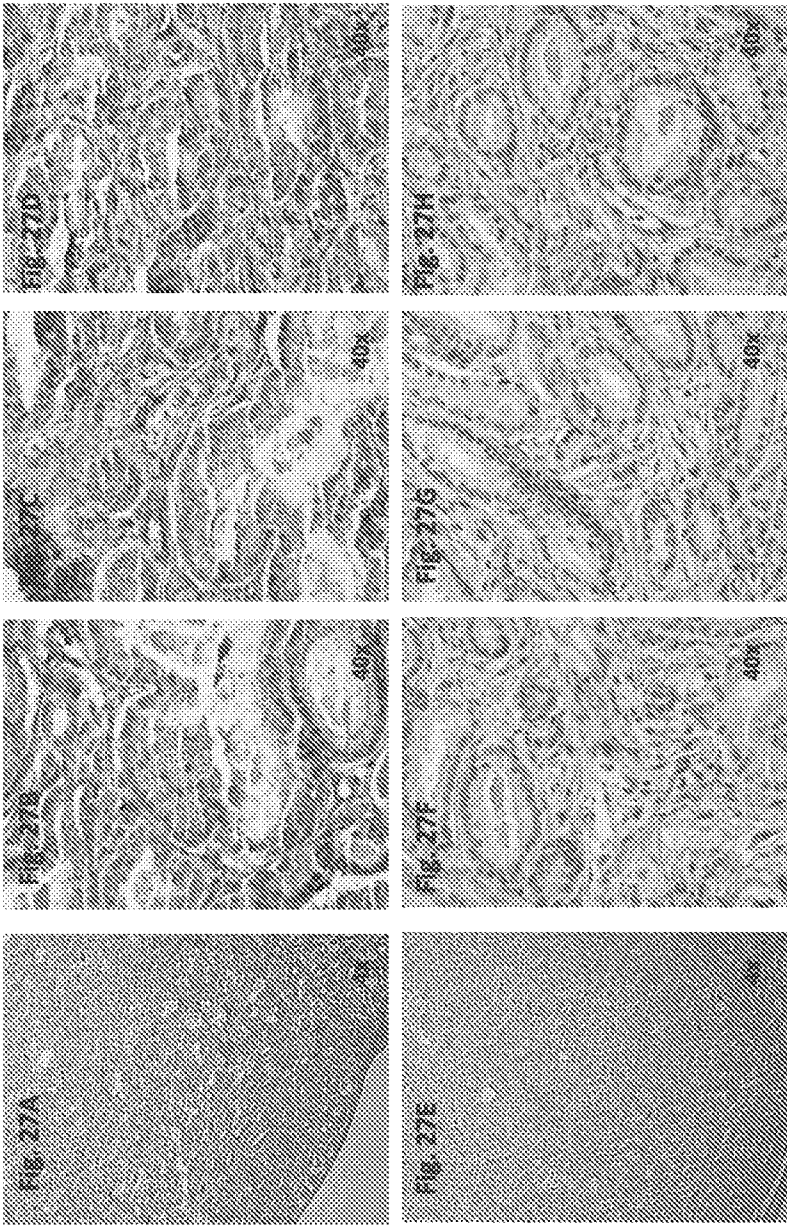
50 ug/ml  
2' (1:1000)

2' only control  
(GaH-HRP)  
1:1000

Figure 26A-26H



Prostate Cancer  
HUCAT361  
Grade 1; Gleason 2; 2+2



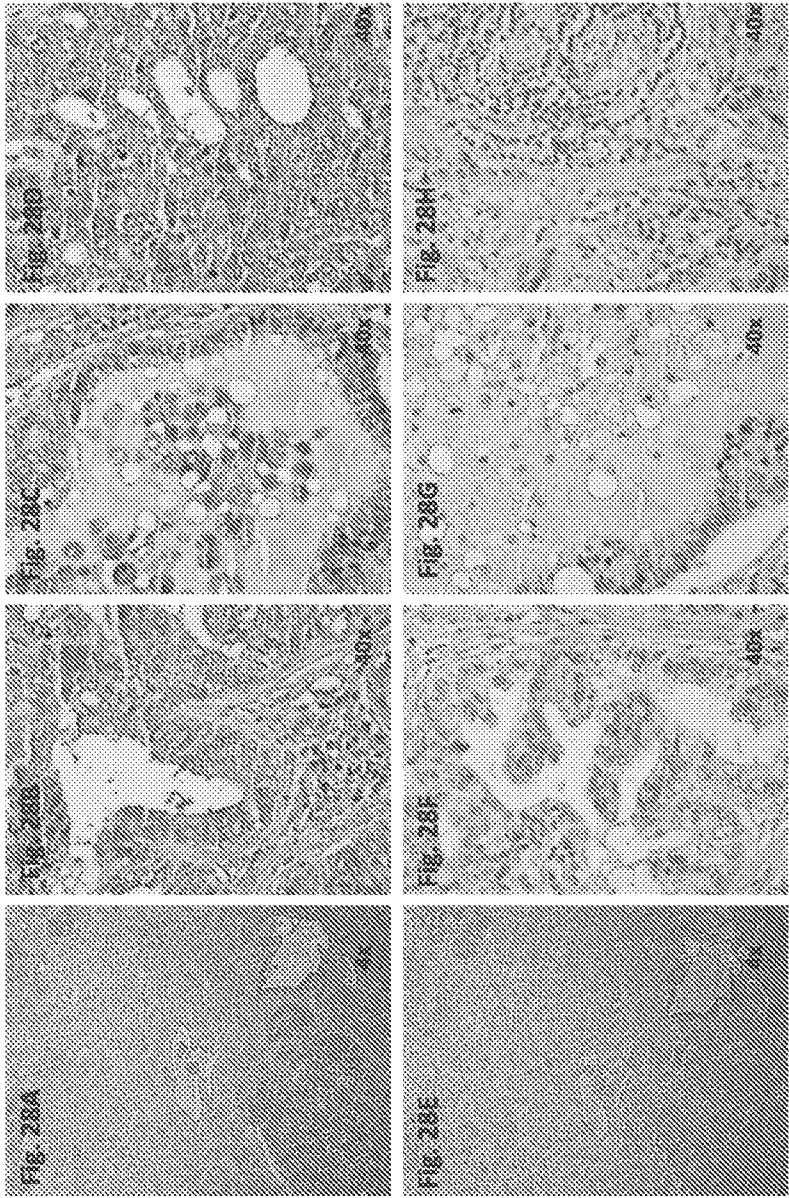
hu-E6  
scFv+Fc

50 ug/ml  
2' (1:750)

2' only control  
(GaH-HRP)  
1:750

Figure 27A-27H

Prostate Cancer  
HUCAT370  
Grade 3; Gleason 3; 3+4

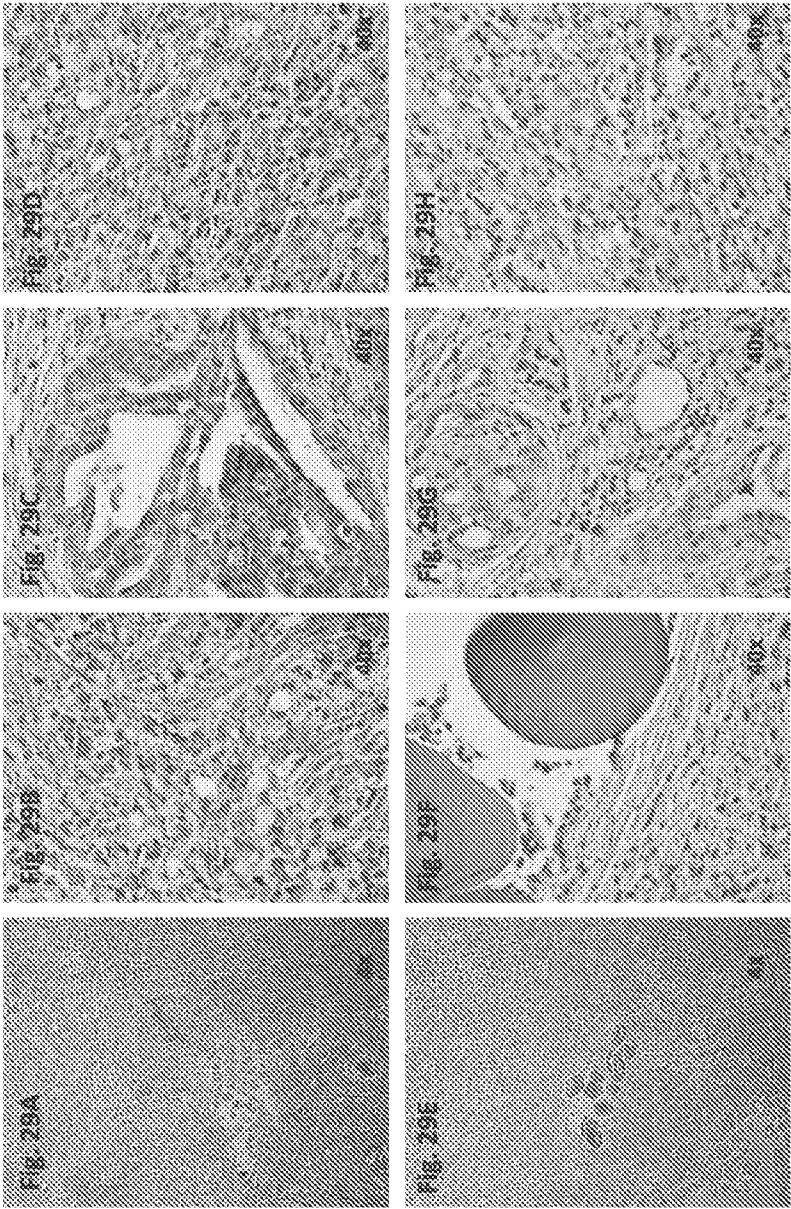


hu-E6  
scFv+Fc  
50 ug/ml  
2' (1:750)

2' only control  
(GaH-HRP)  
1:750

Figure 28A-28H

Prostate Cancer  
HUCAT362  
Grade 4; Gleason 5; 5+5



hu-E6  
scFv+Fc

50 ug/ml  
2' (1:750)

2' only control  
(GalH-HRP)  
1:750

Figure 29A-29H



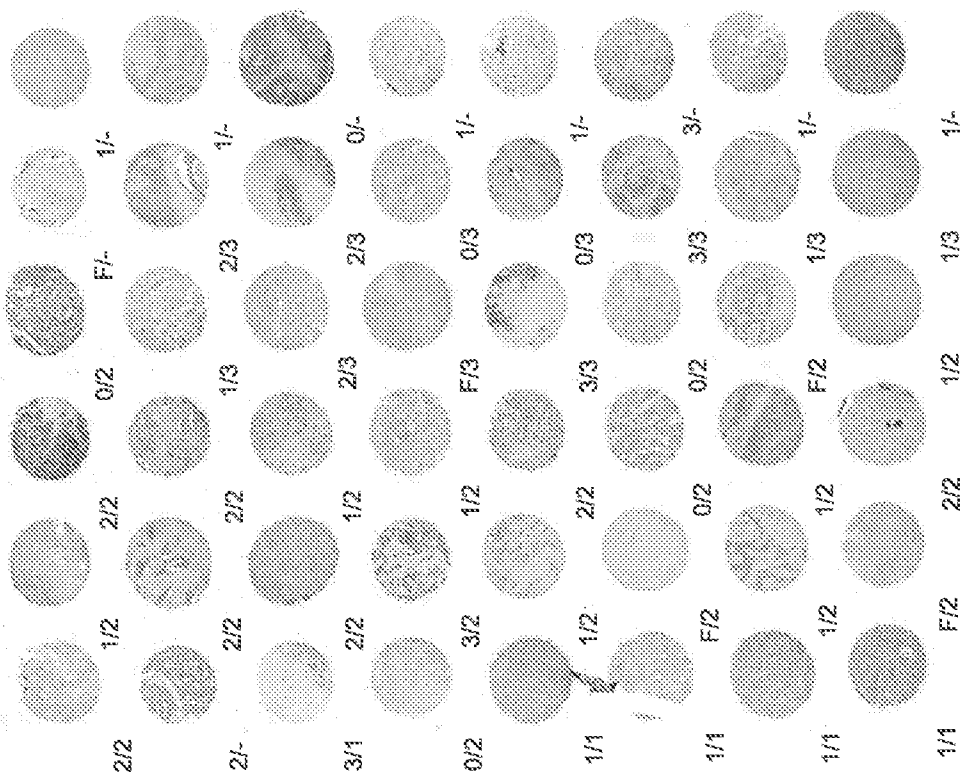
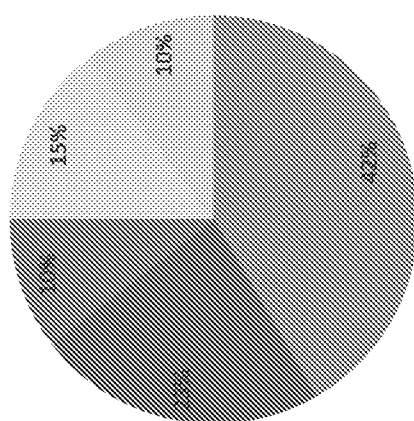


Figure 30A-30B

**Triple Negative Breast Cancer 85% Positive**

32437

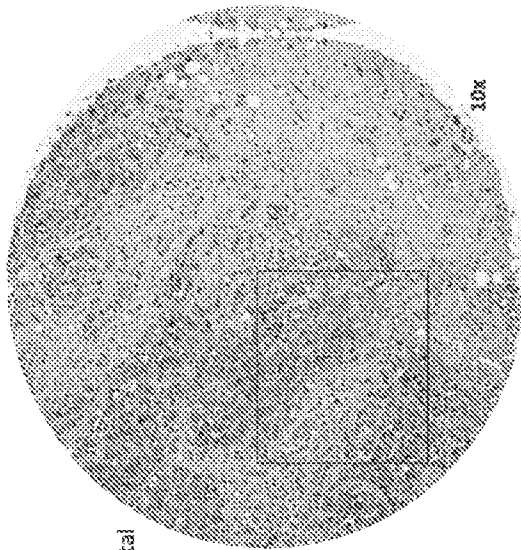
Antibody: MN-hC2-scFv-Fc-Biotin 50 $\mu$ g/ml

300

■ Negative ■ Focal ■ Score 1 ■ Score 2 ■ Score 3

Score/Grade

2/3 = Score 2; Grade 3



Score 3  
Grade 3  
T2N1M0  
Invasive ductal  
Carcinoma  
Address F3

Fig. 30D

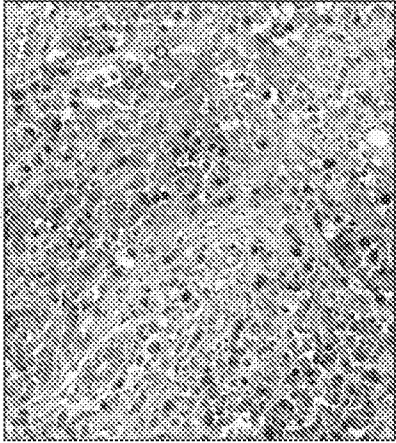
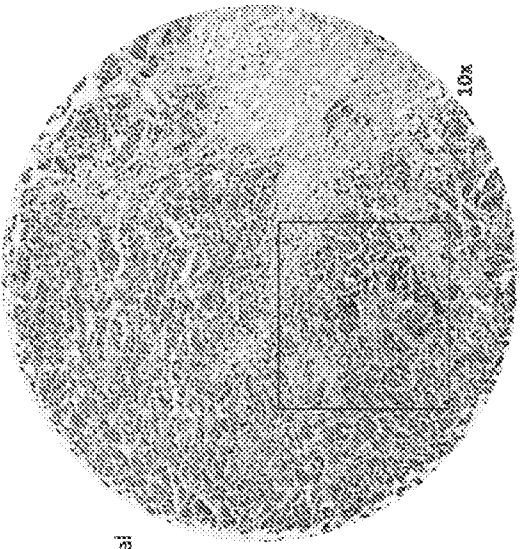


Fig. 30F



Score 2  
Grade 2  
T2N0M0  
Invasive ductal  
Carcinoma  
Address C8

Fig. 30C

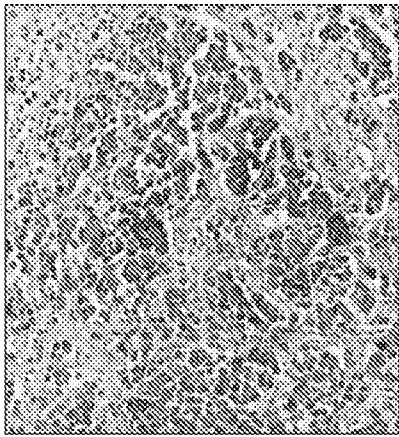
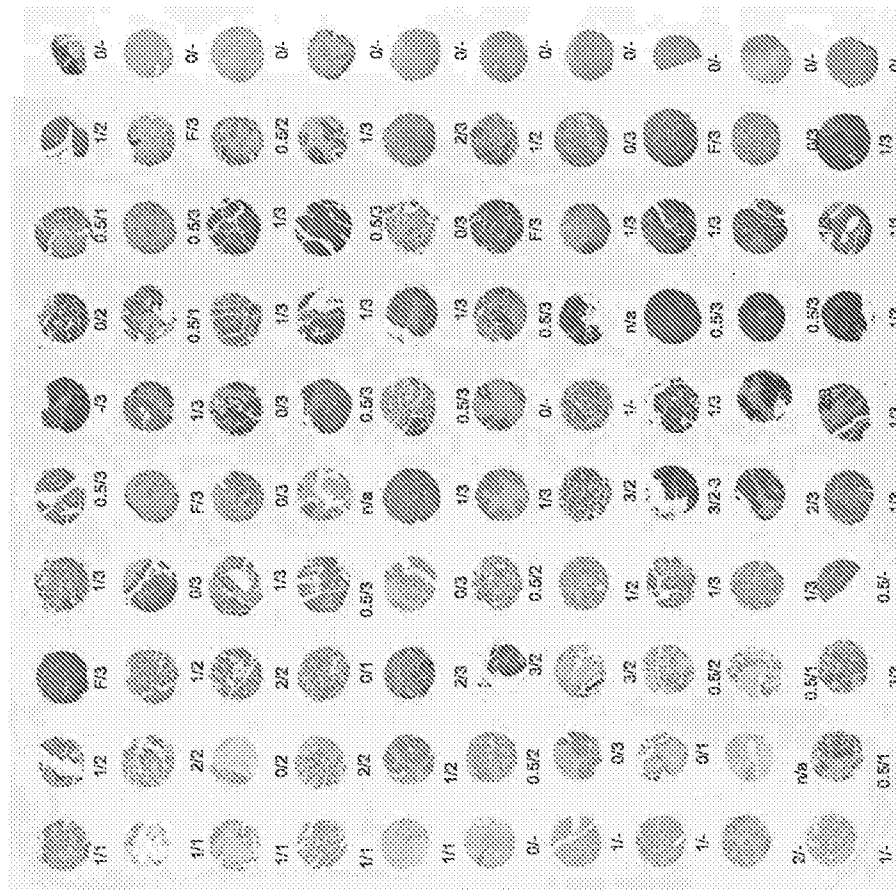


Fig. 30E

Figure 30C-30F



**Ovarian Cancer 83% Positive**  
Biomax BC1115a  
Antibody: MN-hC2-scFv-Fc-Biotin 50ug/mL

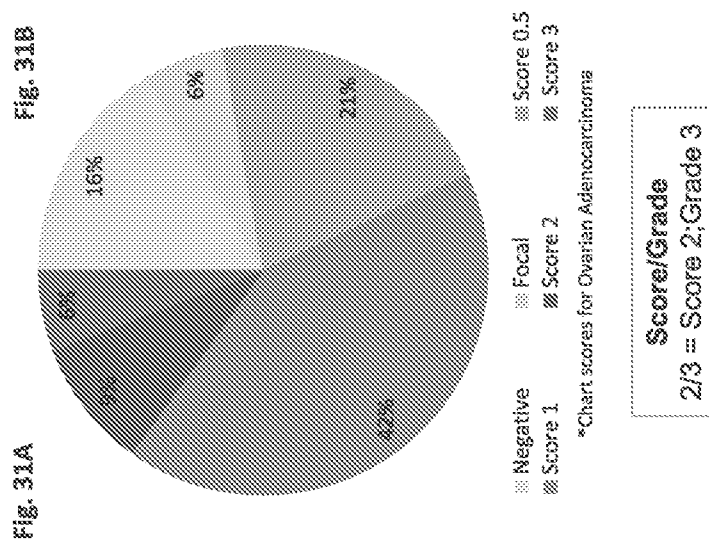
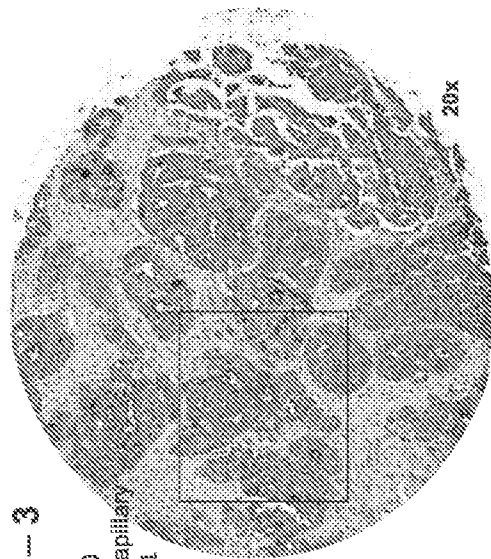


Figure 34-318



Score -- 3  
Grade 3  
T2bNOM0  
Serous Papillary  
Address C1

Fig. 31D

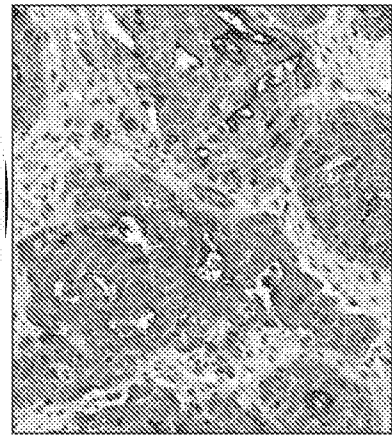
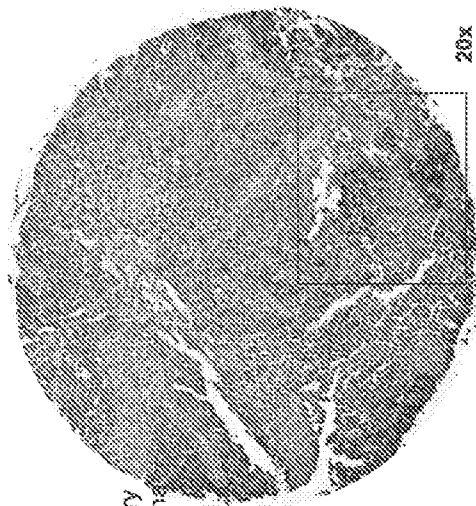


Fig. 31F



Score 2  
Grade 3  
T1aNOM0  
Serous Papillary  
Adenocarcinoma  
Address C6

Fig. 31C

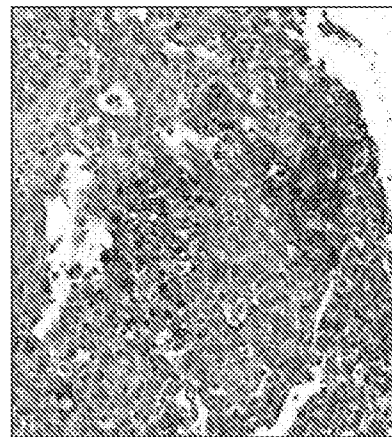


Fig. 31E

Figure 31C-31F

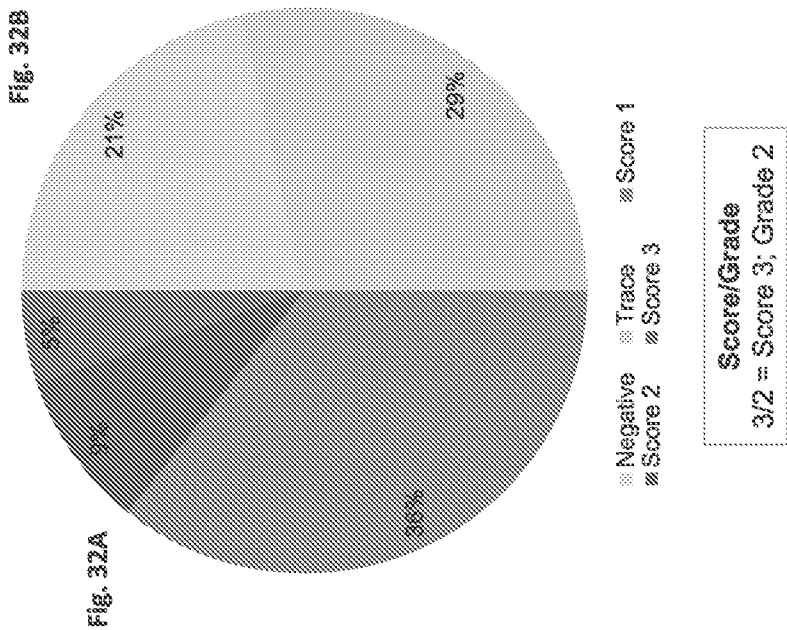
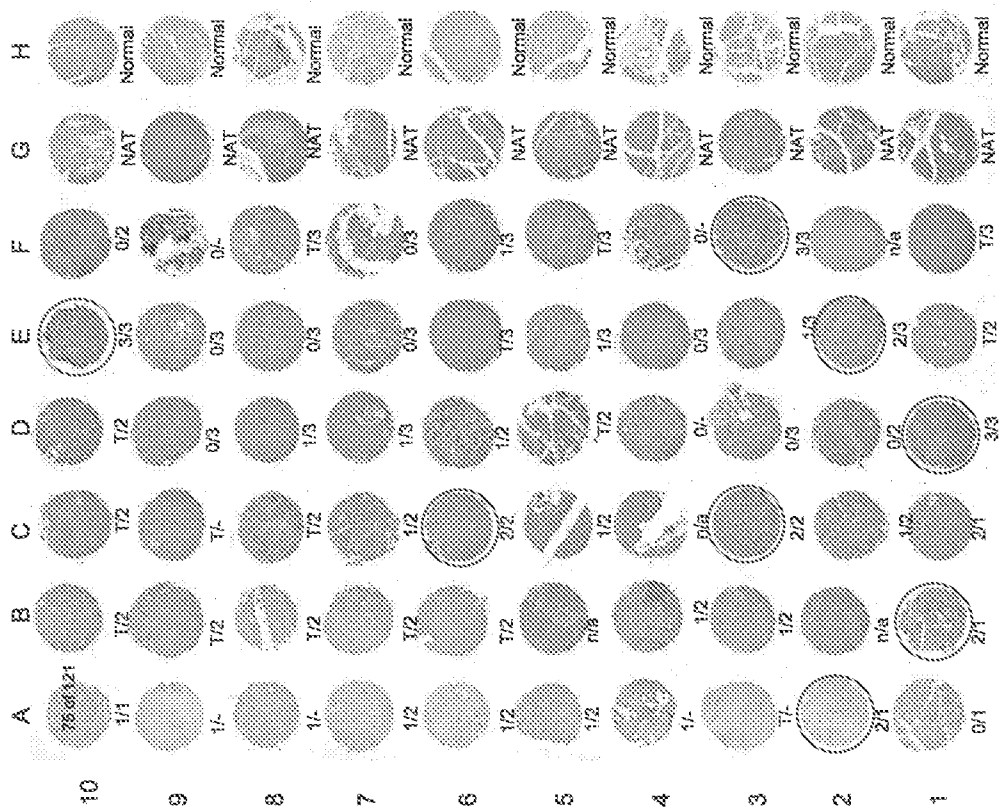


Figure 32A-32B

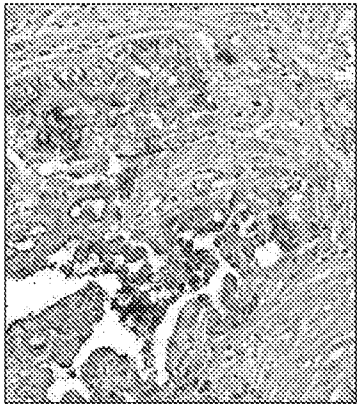
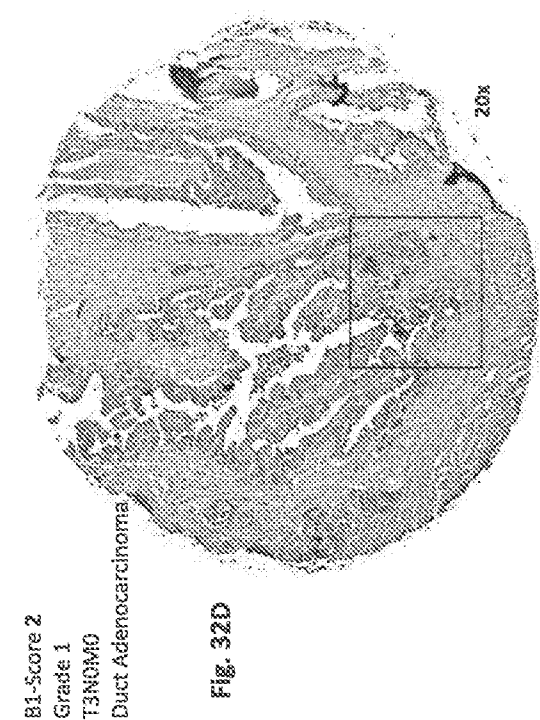


Fig. 32F

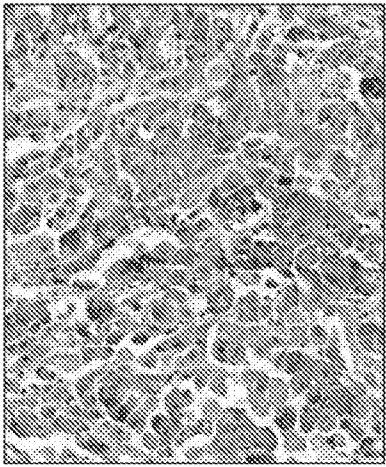
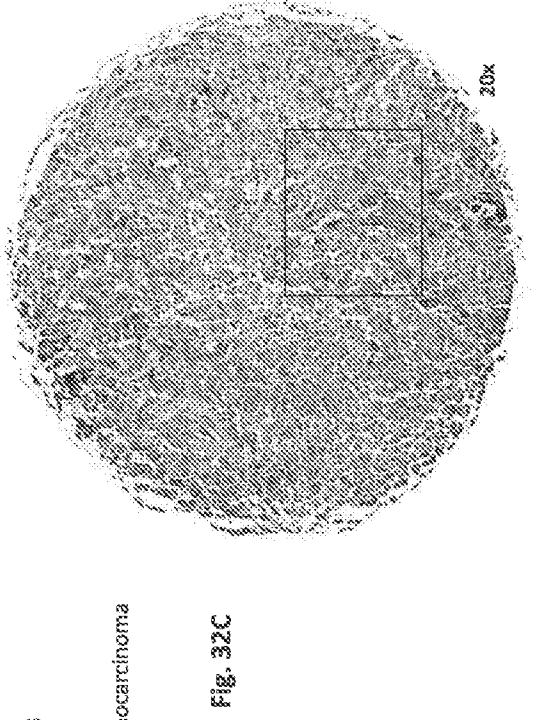
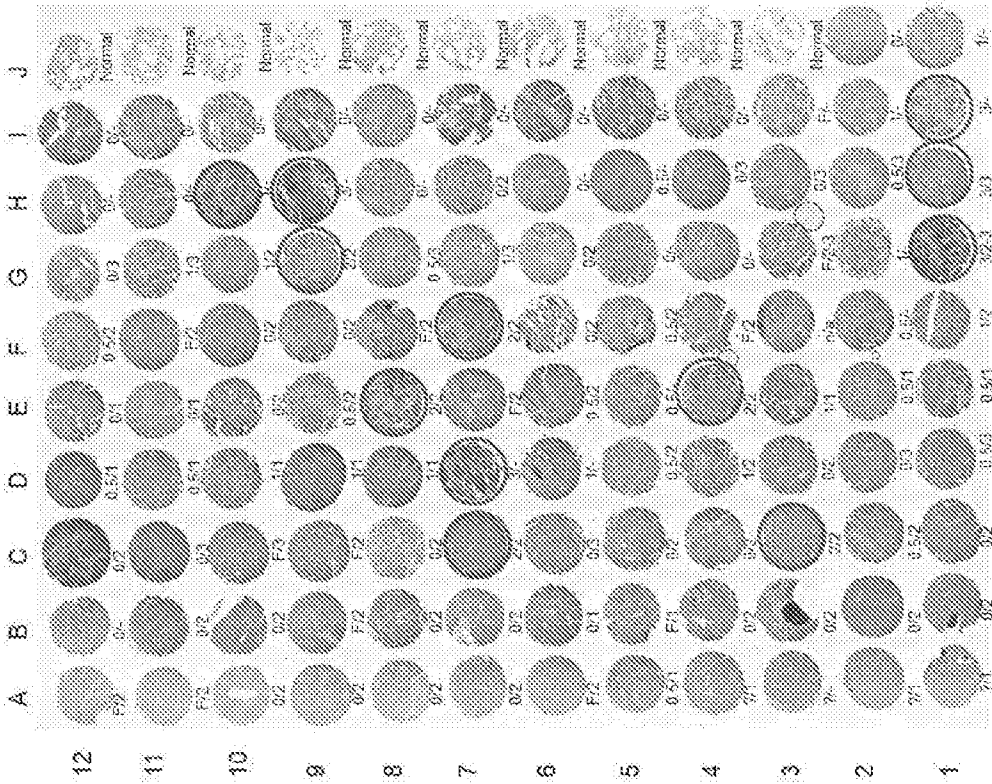


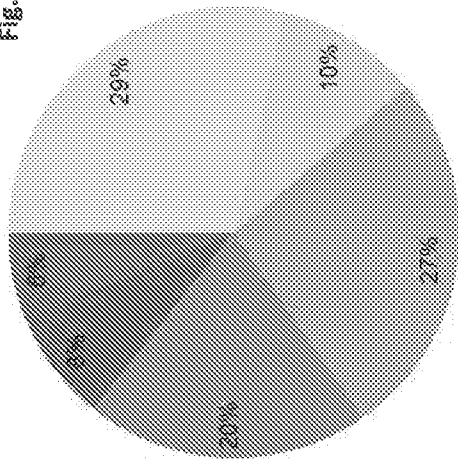
Fig. 32E

Figure 32C-32F



**Lung Cancer Array**  
Antibody: MN-hC2-scFv-Fc-Biotin 50ug/mL

Fig. 33B



■ Negative   ■ Focal   ■ Score 0.5  
■ Score 1   ■ Score 2   ■ Score 3

\*Chart scores for Lung Adenocarcinoma

Score/Grade  
3/2 = Score 3; Grade 2

Fig. 33A

Figure 33A-33B



Lung Cancer Array; MN-hC2-scFv-Fc-Biotin [50ug/mL]

I1 - Score 3  
Grade -  
T2N1M0  
Large Cell  
Carcinoma

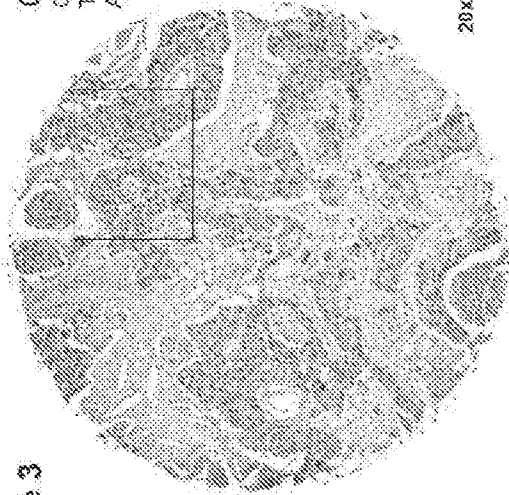


Fig. 33C

G1 - Score 3  
Grade 2-3  
T2N1M0  
Adenocarcinoma

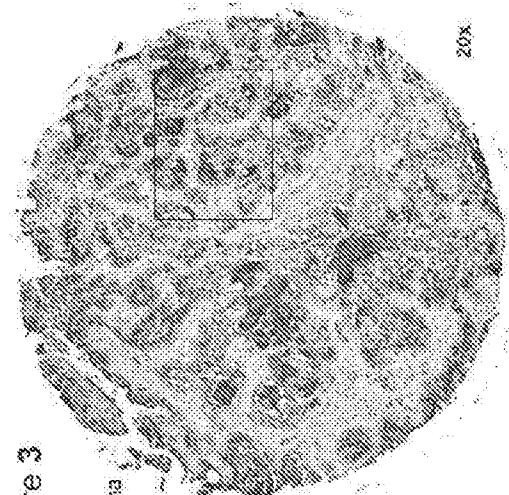


Fig. 33D

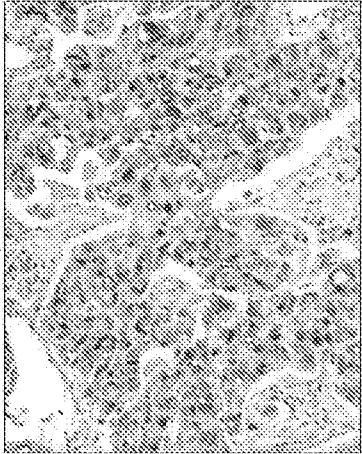


Fig. 33E

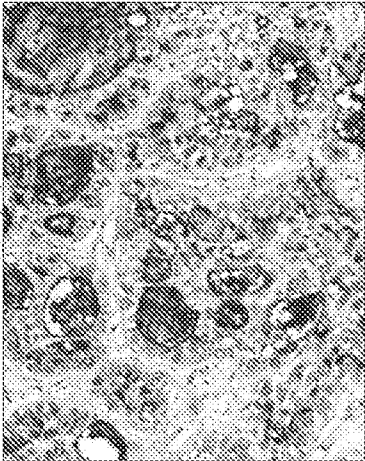


Fig. 33F

Figure 33C-33F



Humanized MNC2 does not bind to full-length MUC1 or MUC1\* on healthy tissues.

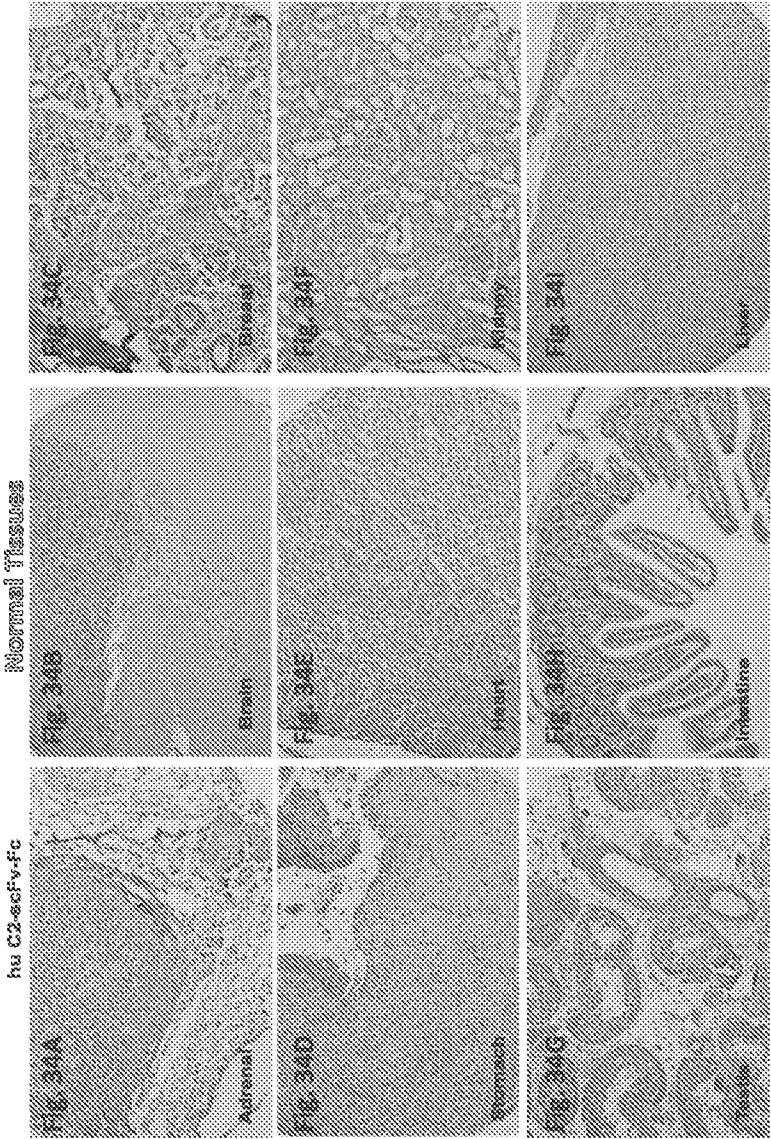


Figure 34A-34I

HCT-116 are a MUC1-negative colon cancer cell line; HCT-MUC1\* is a stable cell line, pre-sorted to be 100% positive for MUC1\*; HCT-MUC1-18 is a single cell clone of HCT's transfected with MUC1-full-length (43 TRs). HCT-MUC1-18 is resistant to MUC1 cleavage (~10% cleaved).

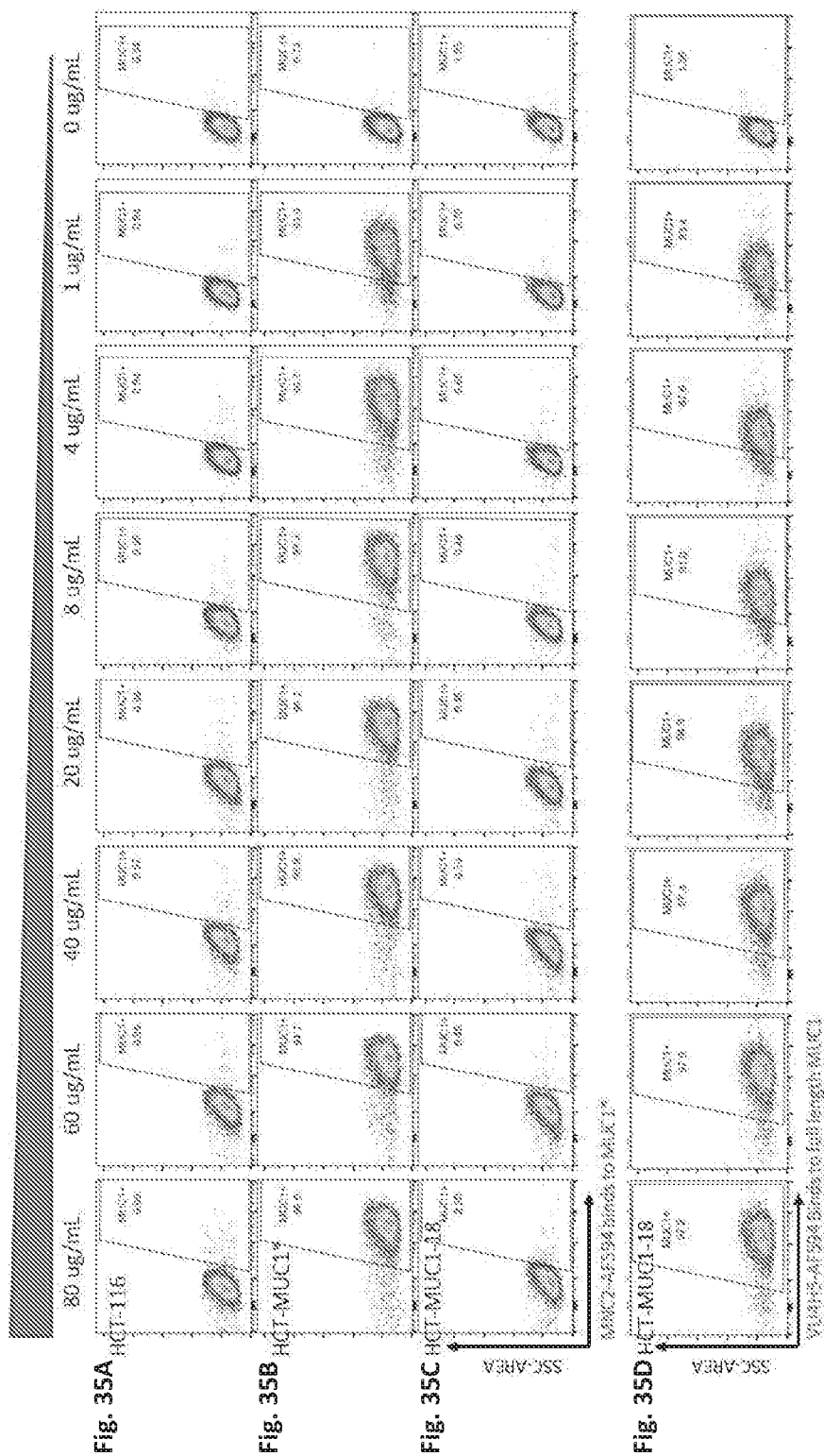


Figure 35A-35D

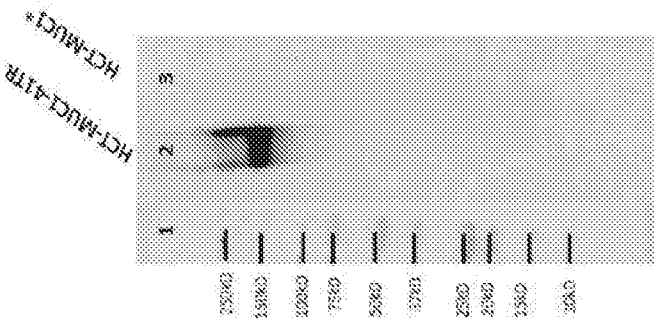


Fig. 36B

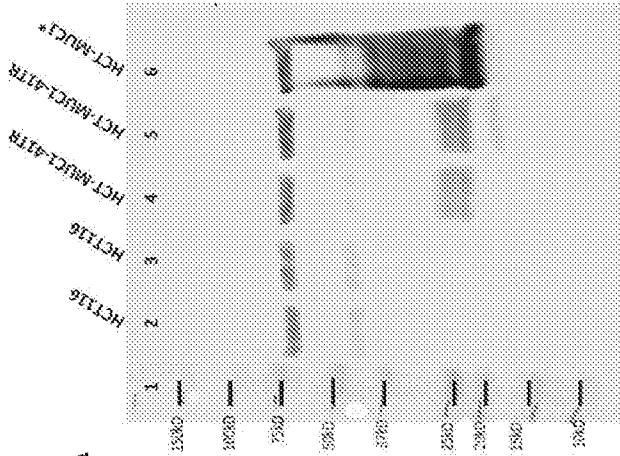


Fig. 36A

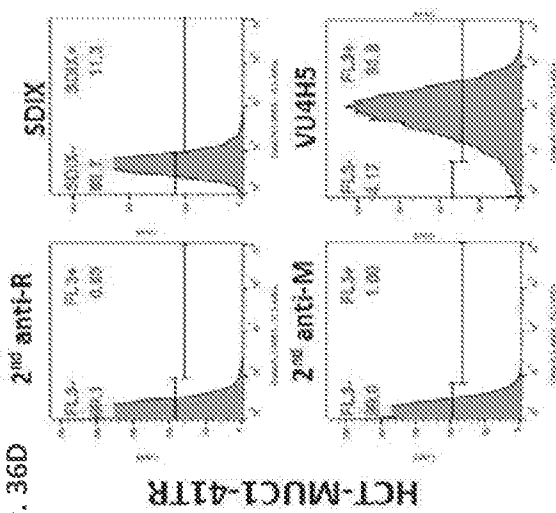


Fig. 36D

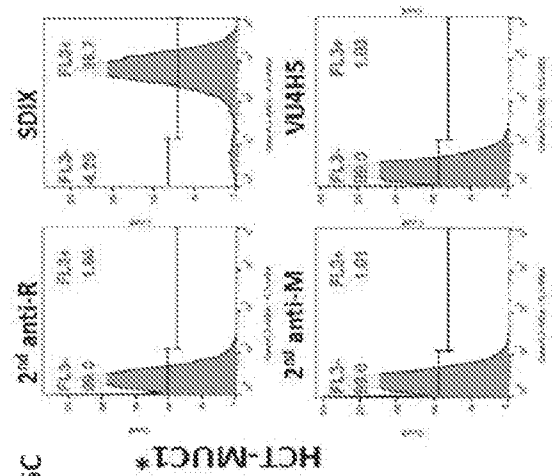


Fig. 36C

Figure 36A-36D

Cleavage releases tandem repeat domain; unmasks binding growth factor sites on MUC1\*

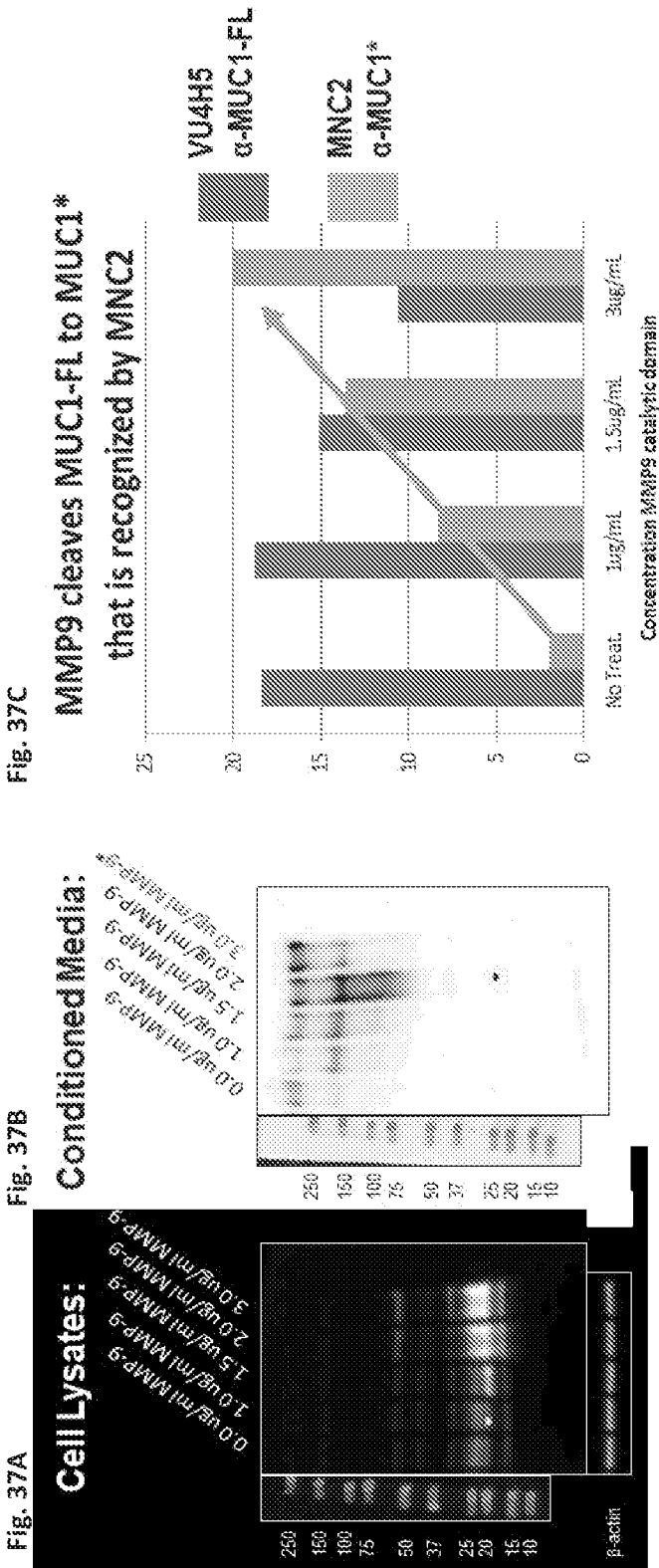


Figure 37A-37C

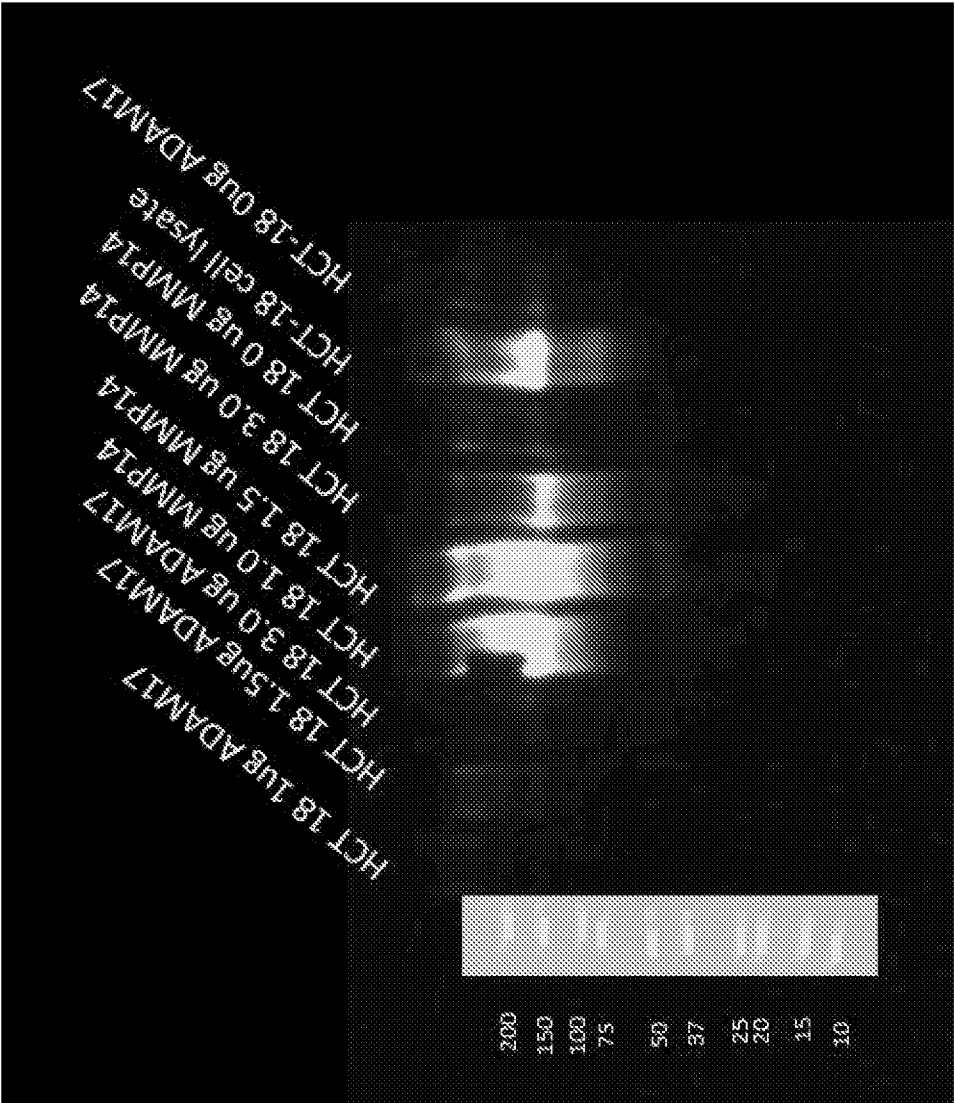


Figure 38

Hematopoietic stem cells express a cleaved MUC1; recognized by SDIX polyclonal & mAb MNC3 but not mAbs MNE6 or MNC2

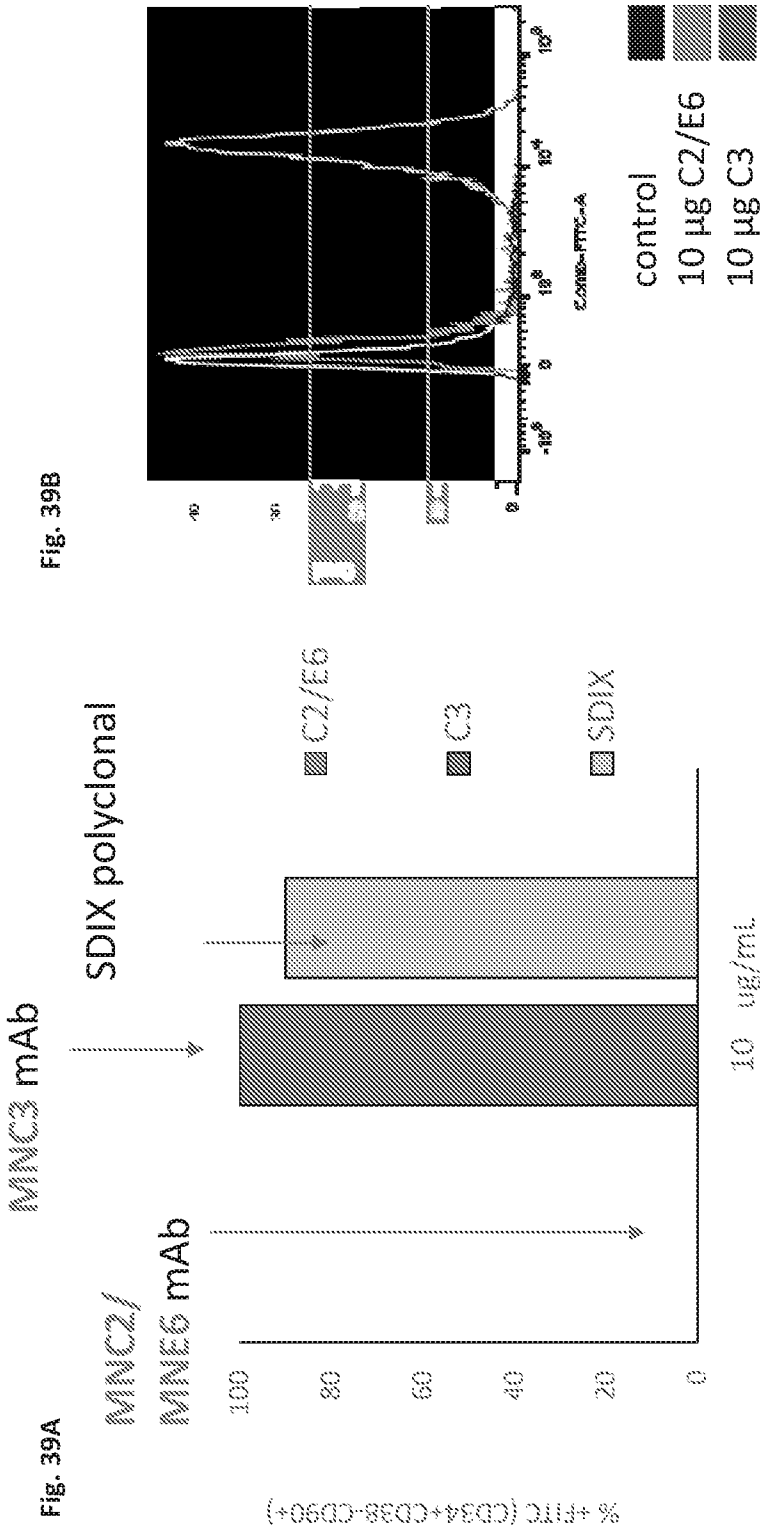


Figure 39A-39B

CD34+/CD38- HSCs are positive for MNC3 but negative for MNE6

Fig. 40A

Total bone marrow cells

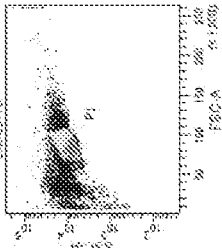


Fig. 40B

CD34+ cells

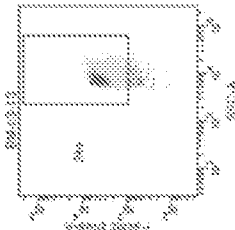


Fig. 40C

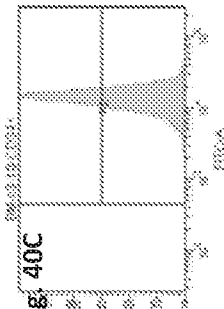


Fig. 40D

CD34+CD38-

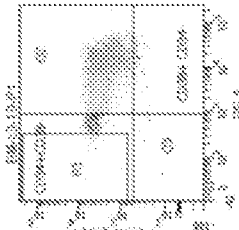


Fig. 40E

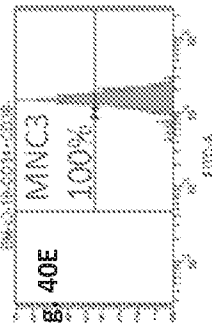


Fig. 40F

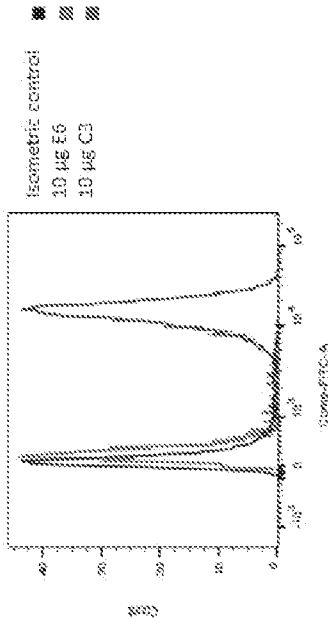


Fig. 40G

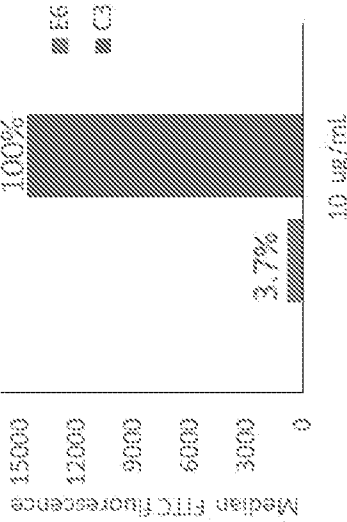


Figure 40A-40G

CD34+/CD38- HSCs stain positive for SDIX anti-MUC1\* polyclonal, but negative for mAbs MNC2 and MNE6

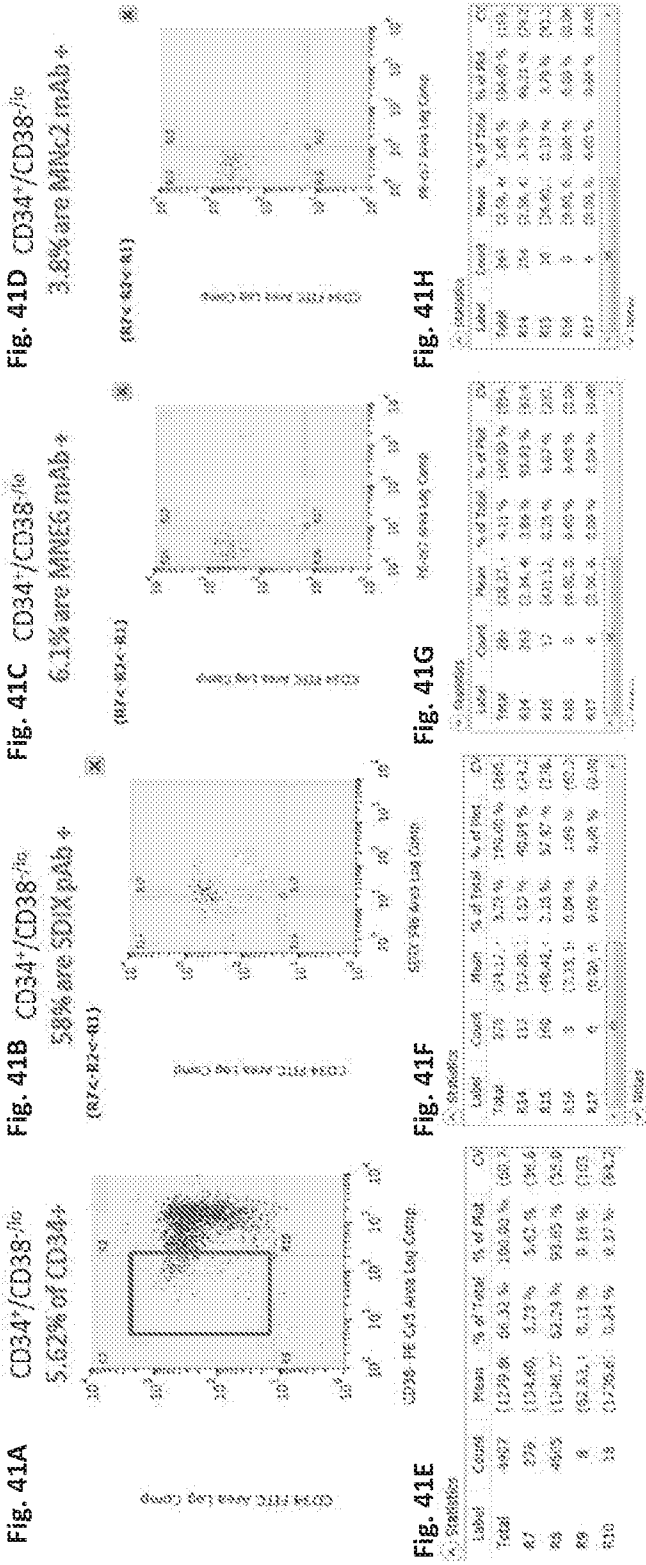


Figure 41A-41H



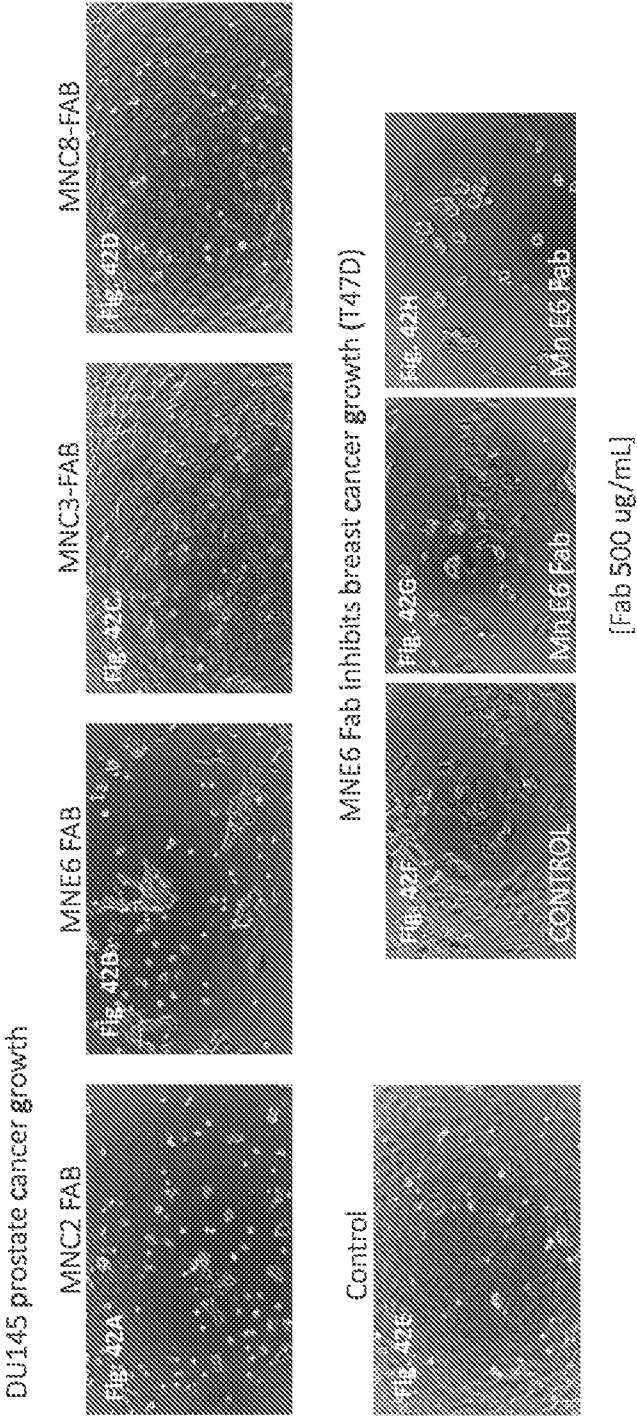


Figure 42A-42H

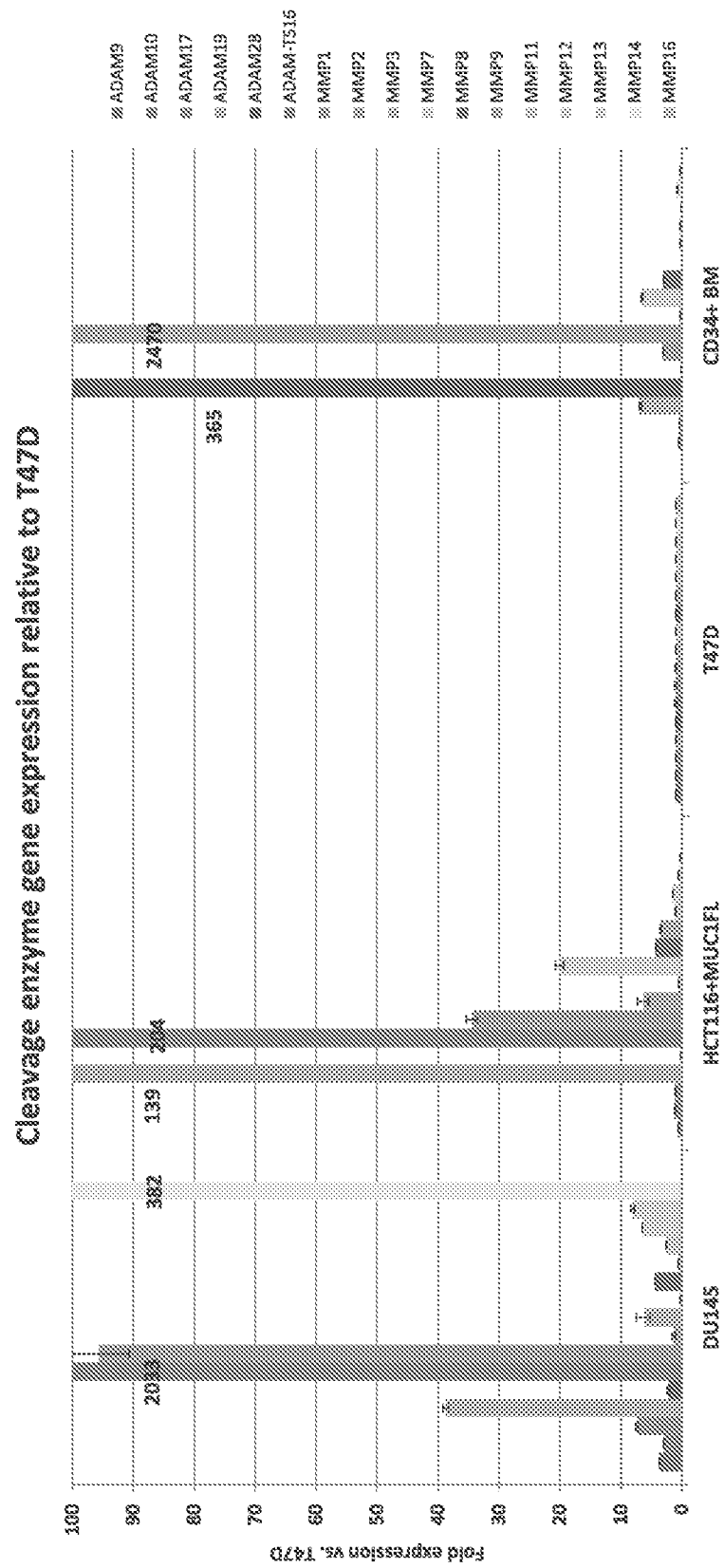


Figure 43

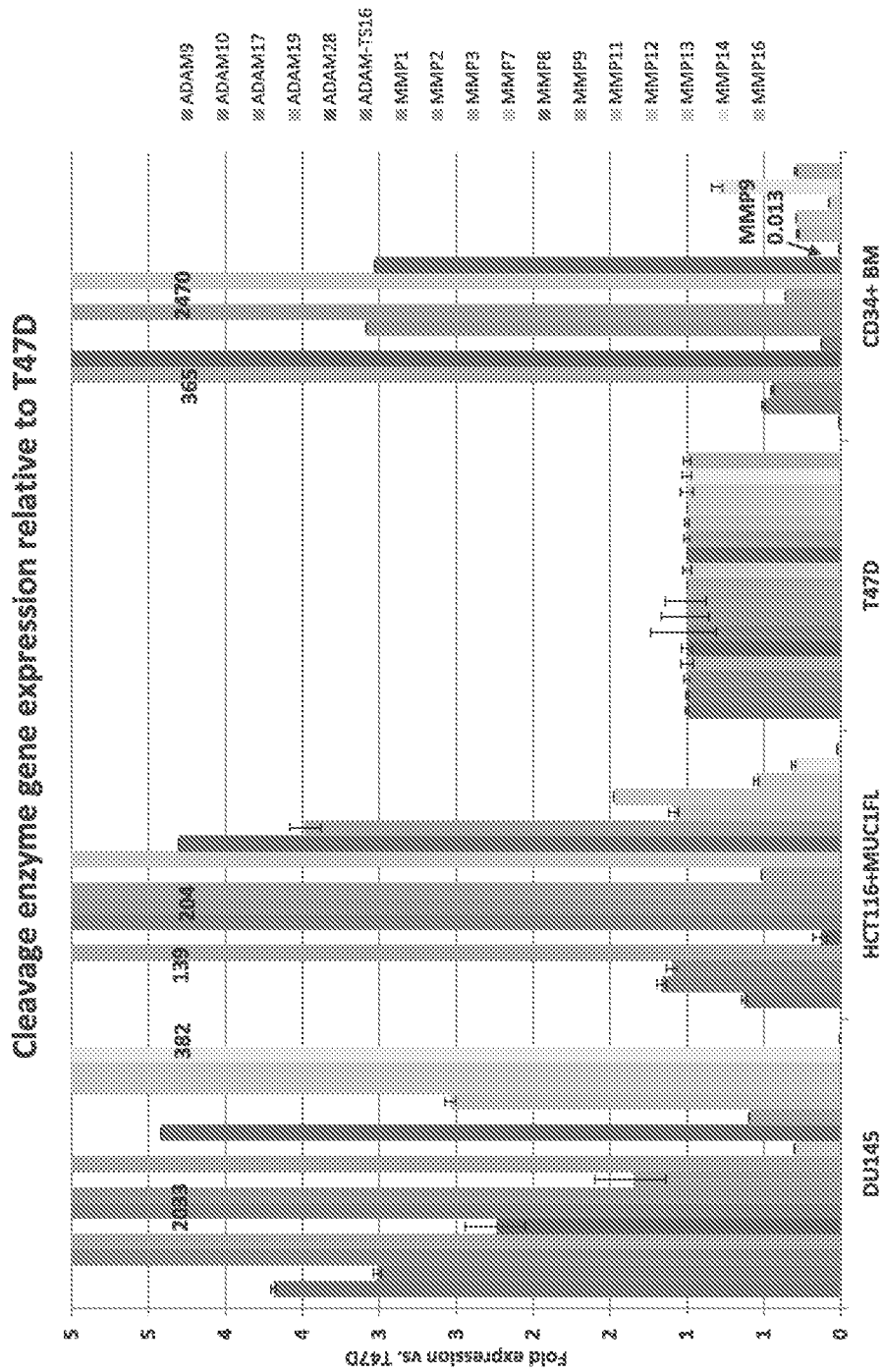


Figure 44

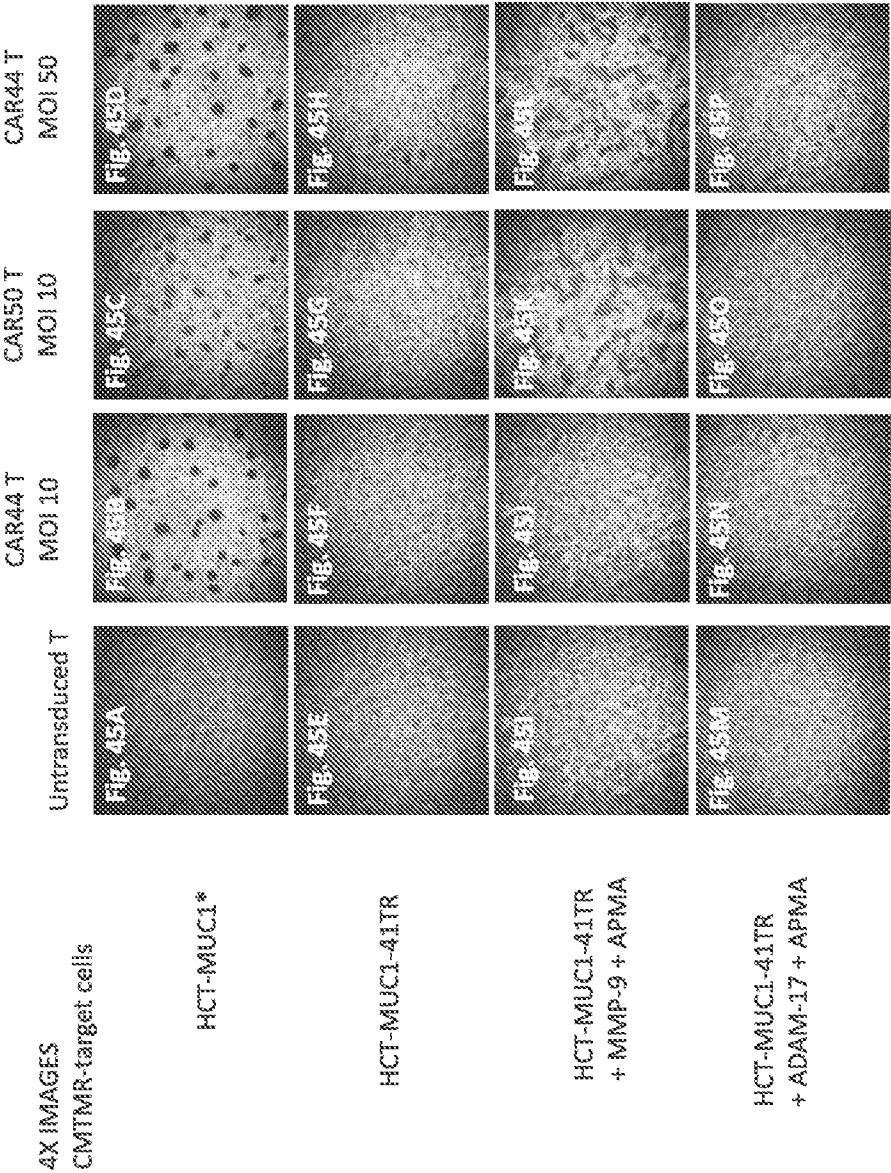


Figure 45A-45P

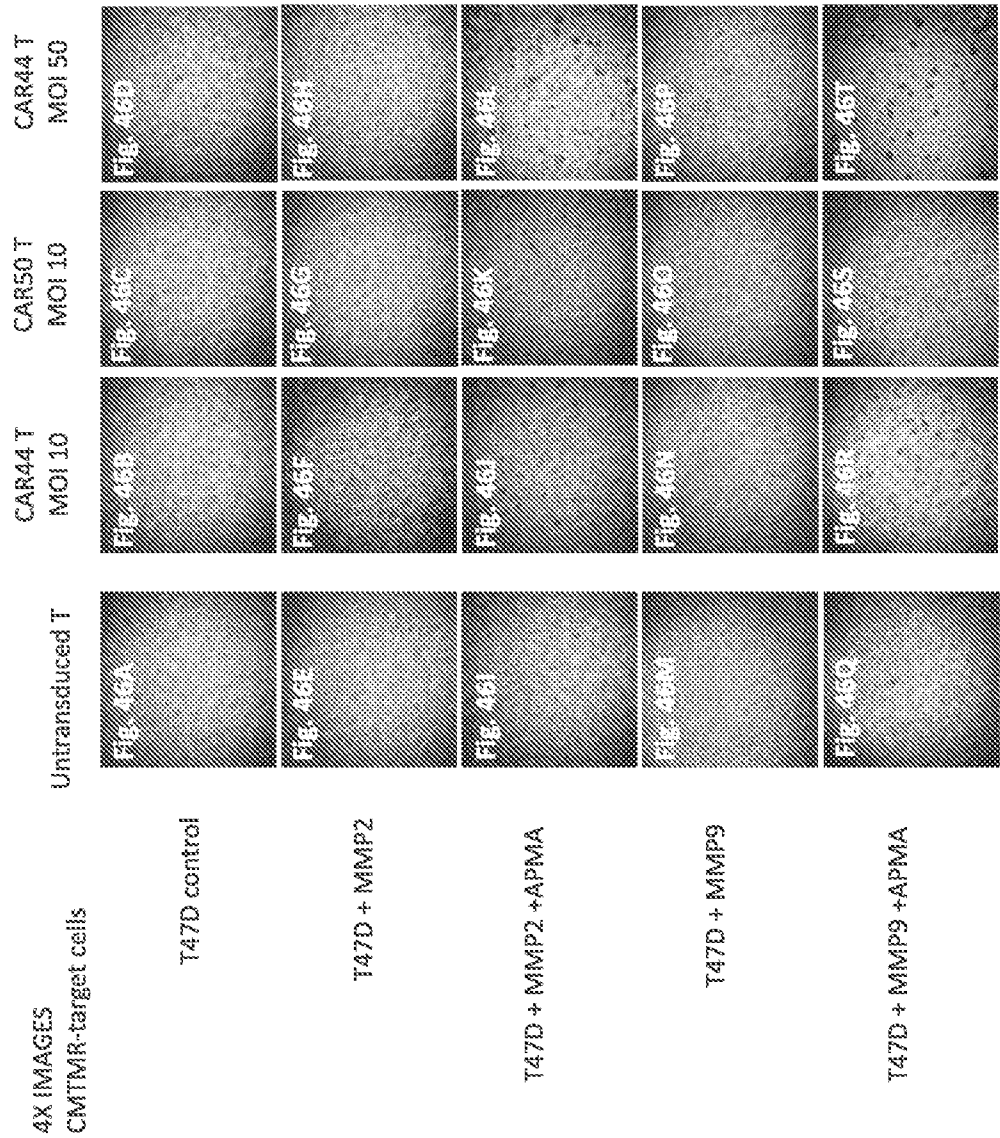


Figure 46A-46T

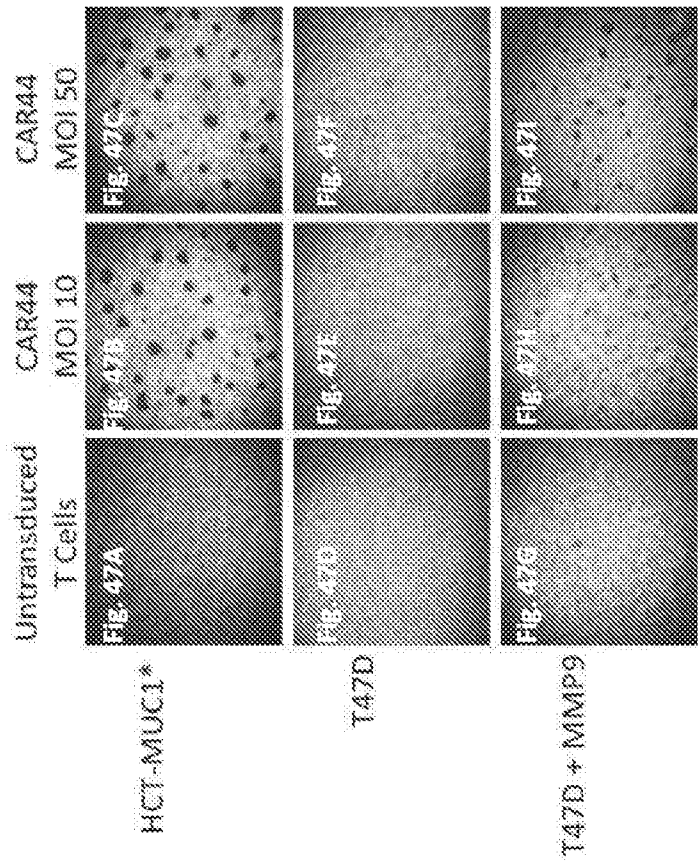


Figure 47A-47I

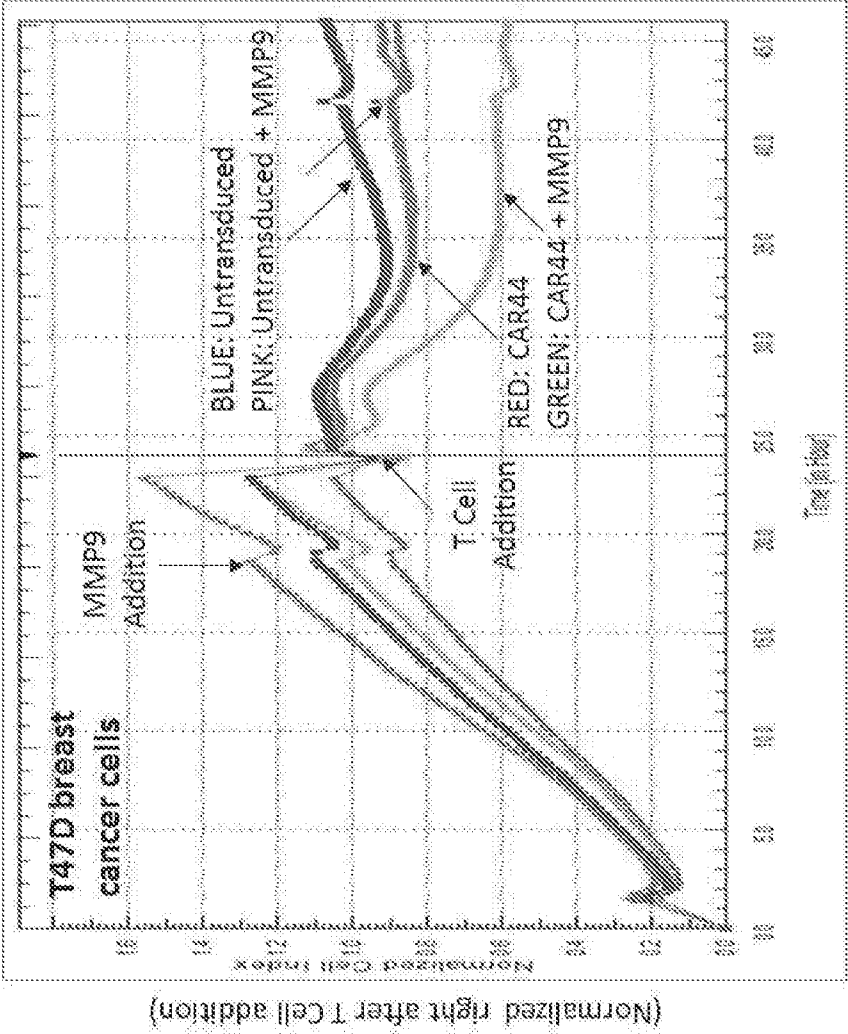


Figure 48

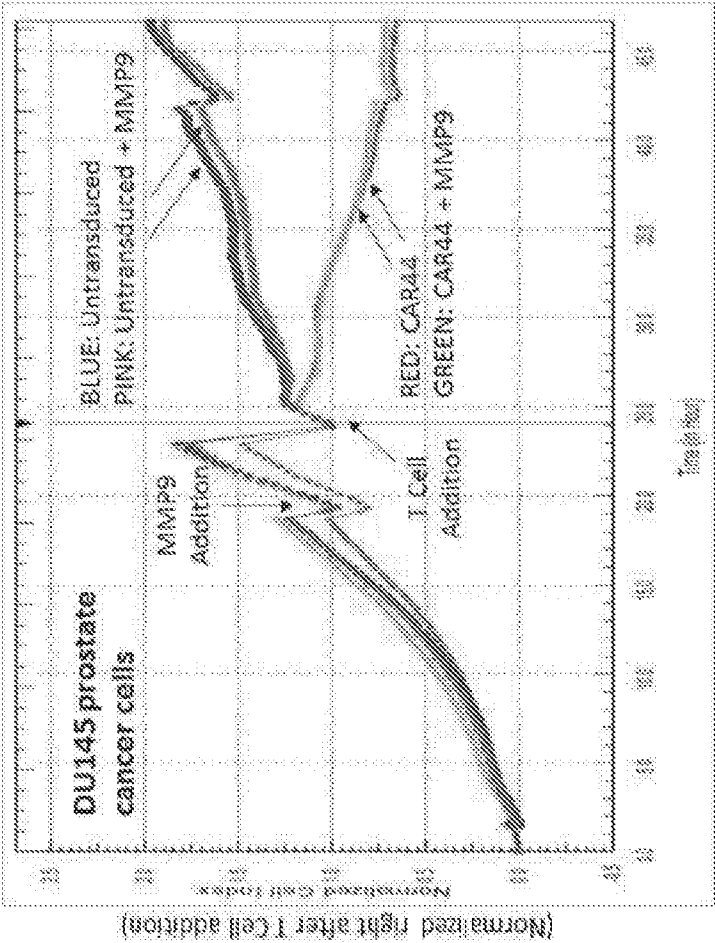


Figure 49



PCR measurement of MUC1 expression in normal cells compared to T47D breast cancer cells and HCT-MUC1\*, a colon cancer cell line transduced with MUC1\*. Measurement was normalized to Hep. G2, MUC1 negative normal liver cells.

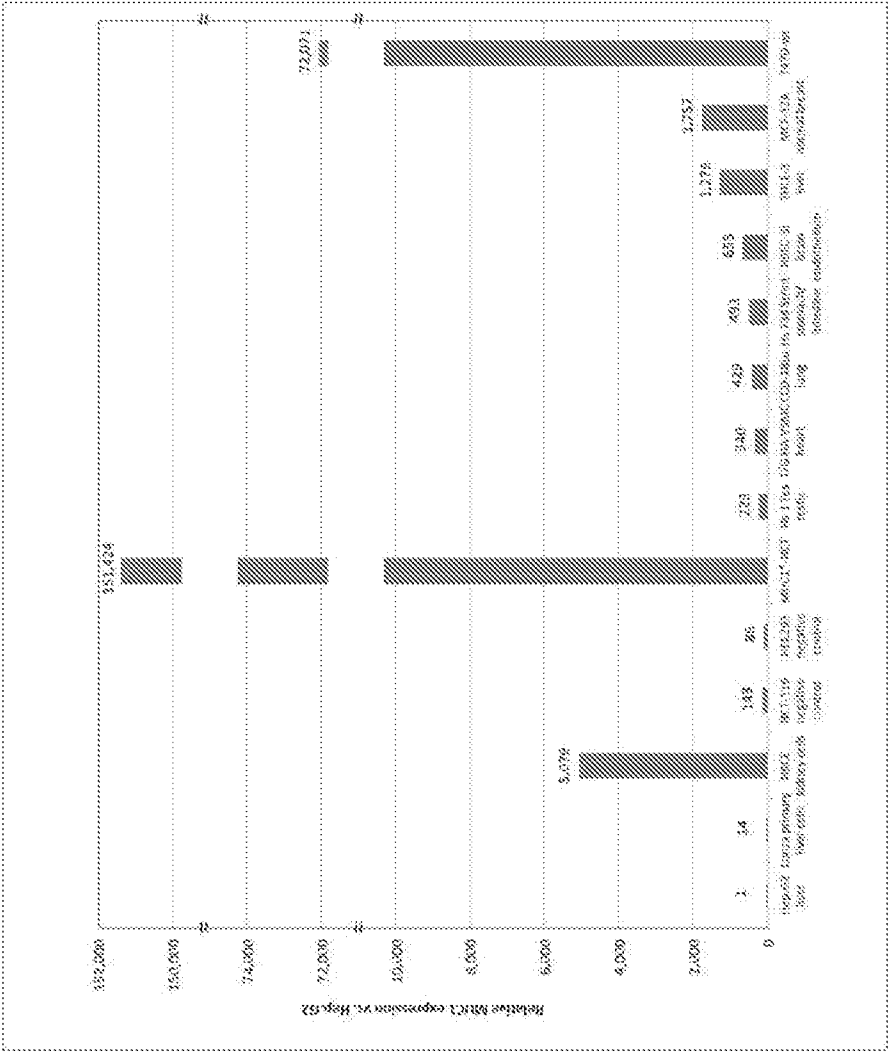


Figure 50

To assess CAR44 T cell activation in response to co-culture with target cells, IFN- $\gamma$  was measured in supernatant of CAR44 T cells in co-culture with normal cells, or cancer cells as a control. IFN- $\gamma$  was measured after 72 hours of co-culture. (A) CAR44 T cell to target cell ratio was 1:1; (B) CAR44 T cell to target cell ratio was 0.5:1.

51A



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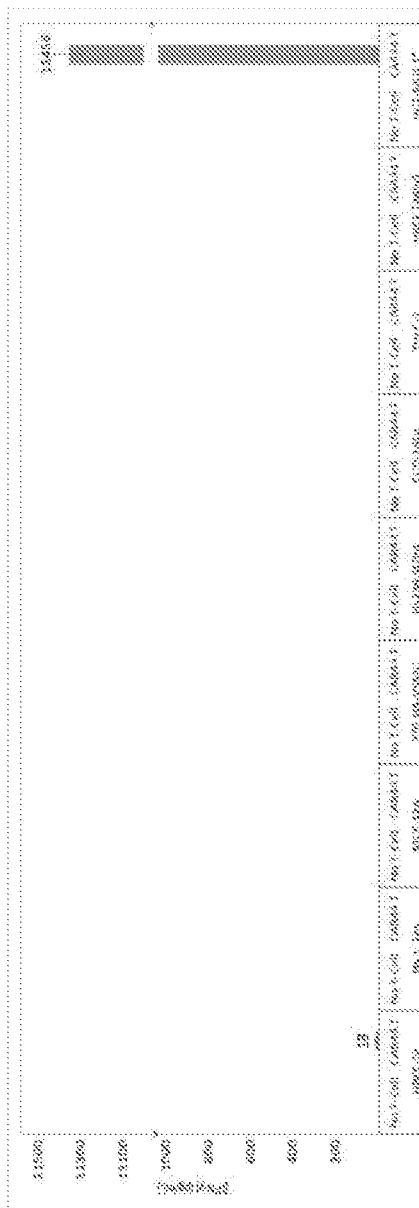


Figure 51A-51B

To assess CAR44 T cell activation in response to co-culture with target cells, IL-2 was measured in supernatant of CAR44 T cells in co-culture with normal cells, or cancer cells as a control. IL-2 was measured after 72 hours of co-culture. (A) CAR44 T cell to target cell ratio was 1:1; (B) CAR44 T cell to target cell ratio was 0.5:1.

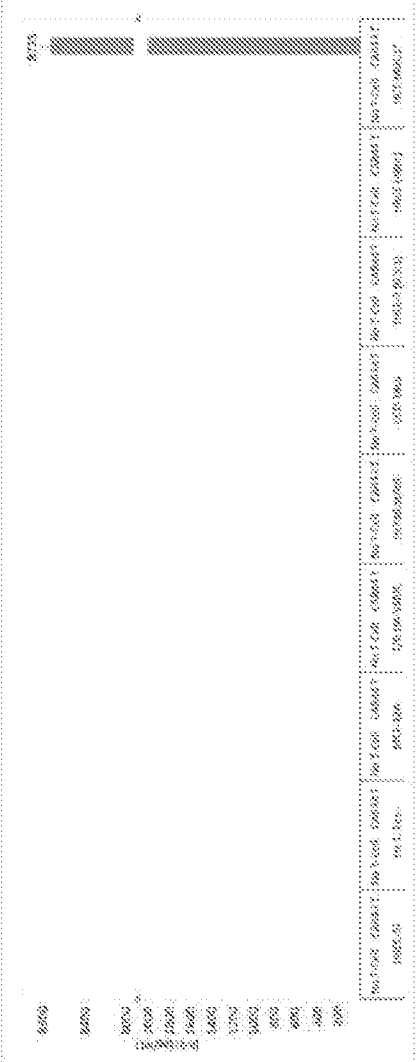
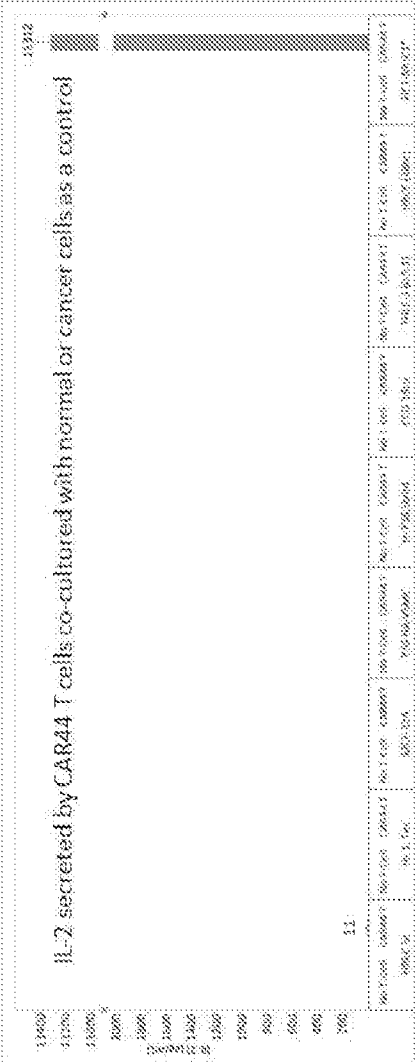


Figure 52A-52B

(1) Normal cells, or cancer cells as a control, were co-culture with no T cells, Untransduced T cells or with CAR44 T cells, at an E:T ratio of 1:1 or 0.5:1. Cells were then labeled with a cell death marker and analyzed by FACS. The CD3+ population (T cells) was eliminated from the cell count. A-F (2) & (3) Magnified photographs of normal cells in co-culture with Untransduced T cells or CAR44 T cells. FACS and magnified photographs were at 48 hours post addition of T cells.

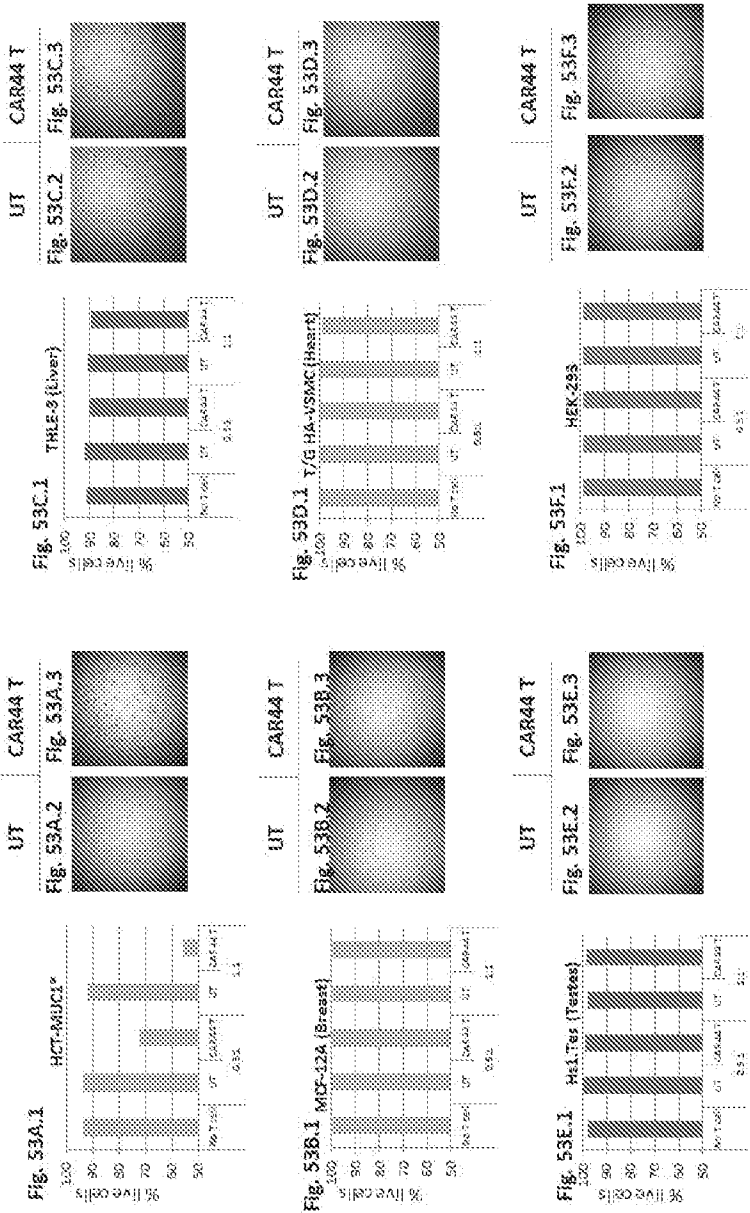


Figure 53A-53F

(1) Normal cells, or cancer cells as a control, were co-culture with no T cells, Untransduced T cells or with CAR44 T cells, at an E:T ratio of 1:1 or 0.5:1. Cells were then labeled with a cell death marker and analyzed by FACS. The CD3+ population (T cells) was eliminated from the cell count. G-J (2) & (3) Magnified photographs of normal cells in co-culture with Untransduced T cells or CAR44 T cells. FACS and magnified photographs were at 48 hours post addition of T cells.

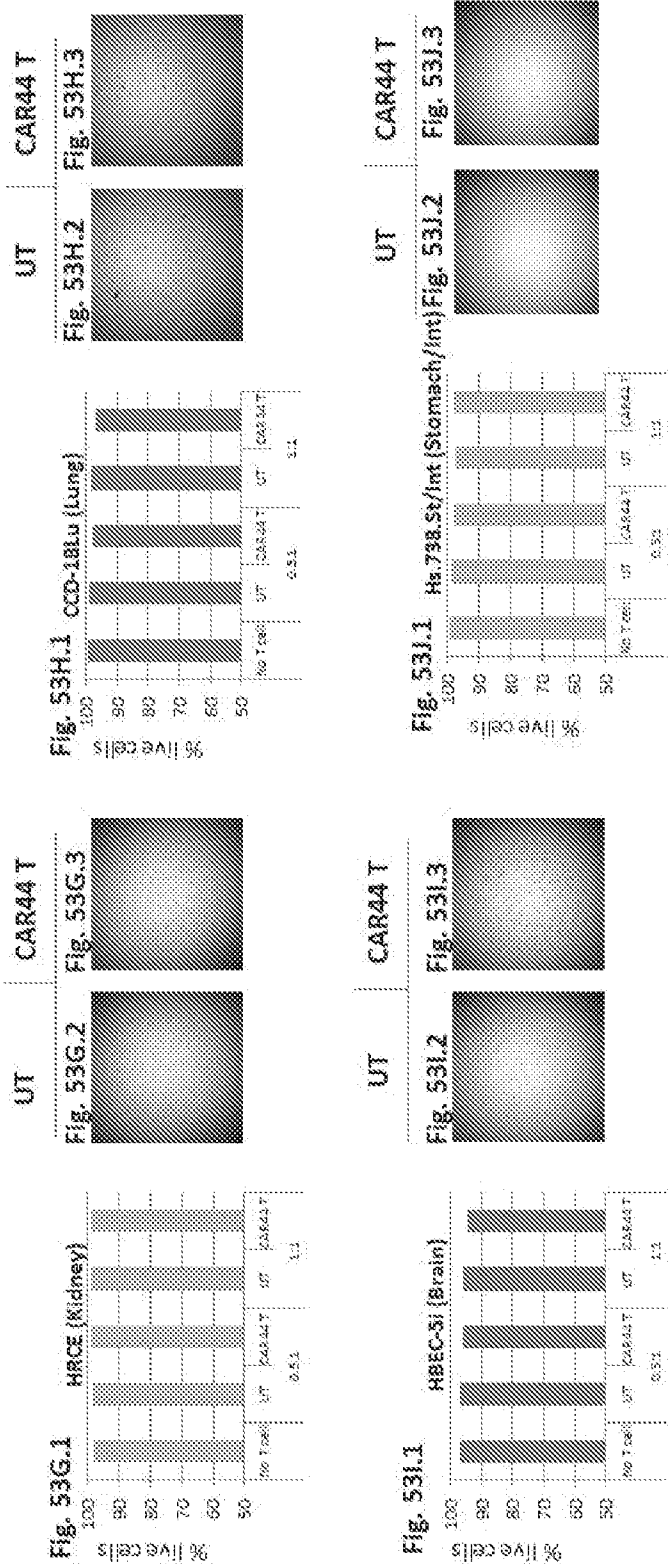


Figure 53G-53J

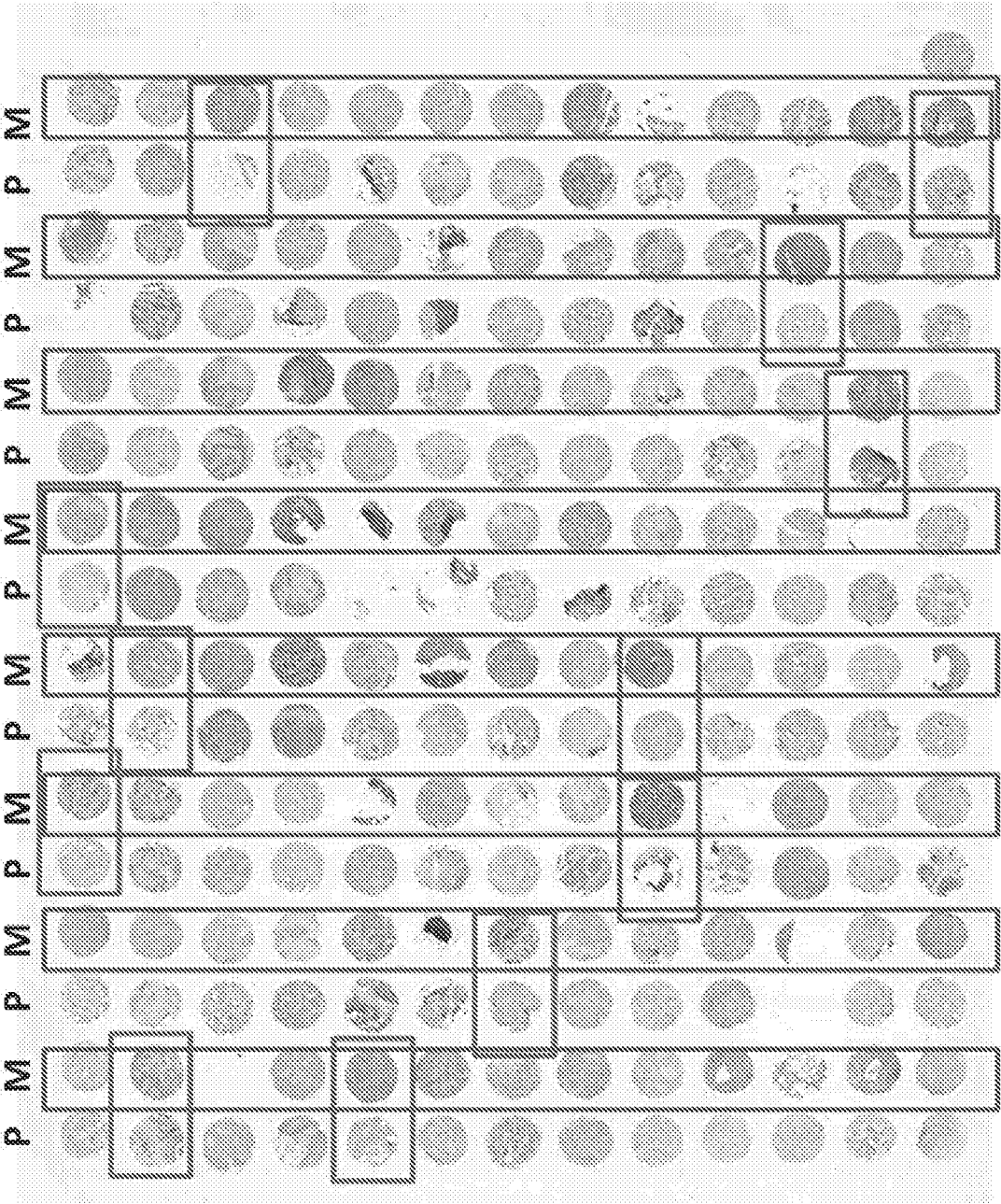


Figure 54

MUC1\* positive prostate cancer cells co-cultured with  
huMNC2-MNCAR44 T cells

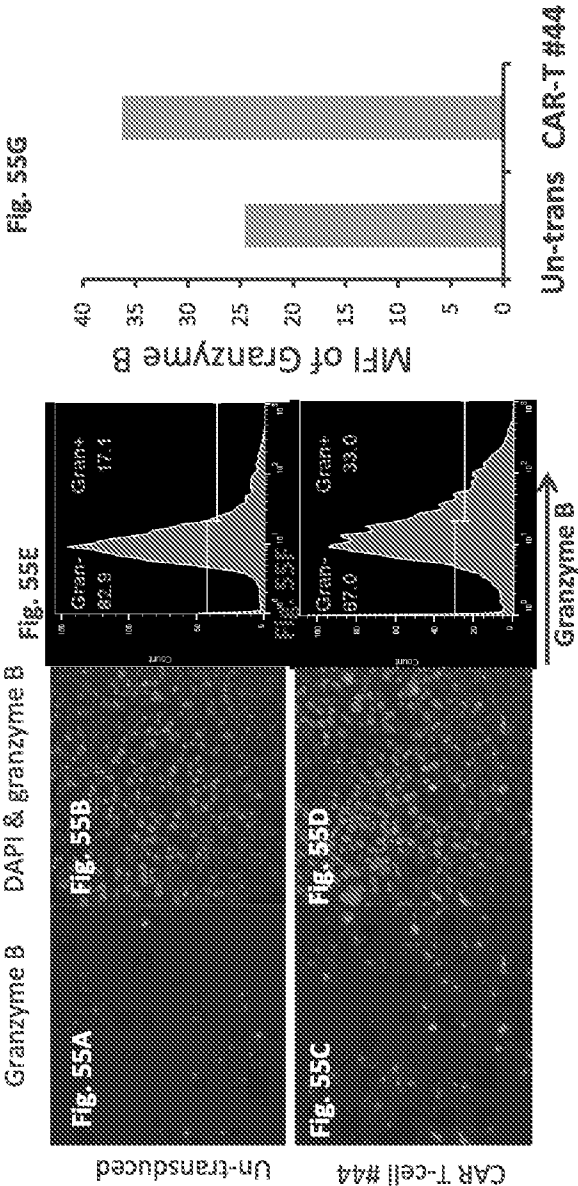


Fig. 55H DU145 MUC1\* positive prostate cancer - 5 days

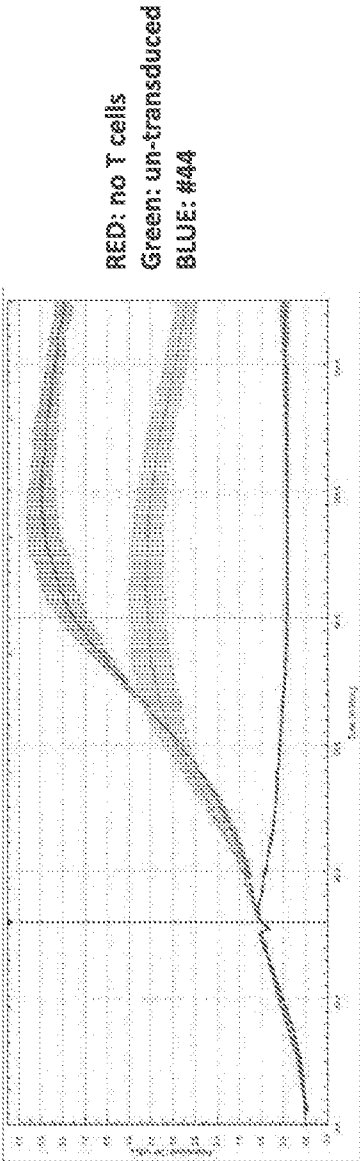
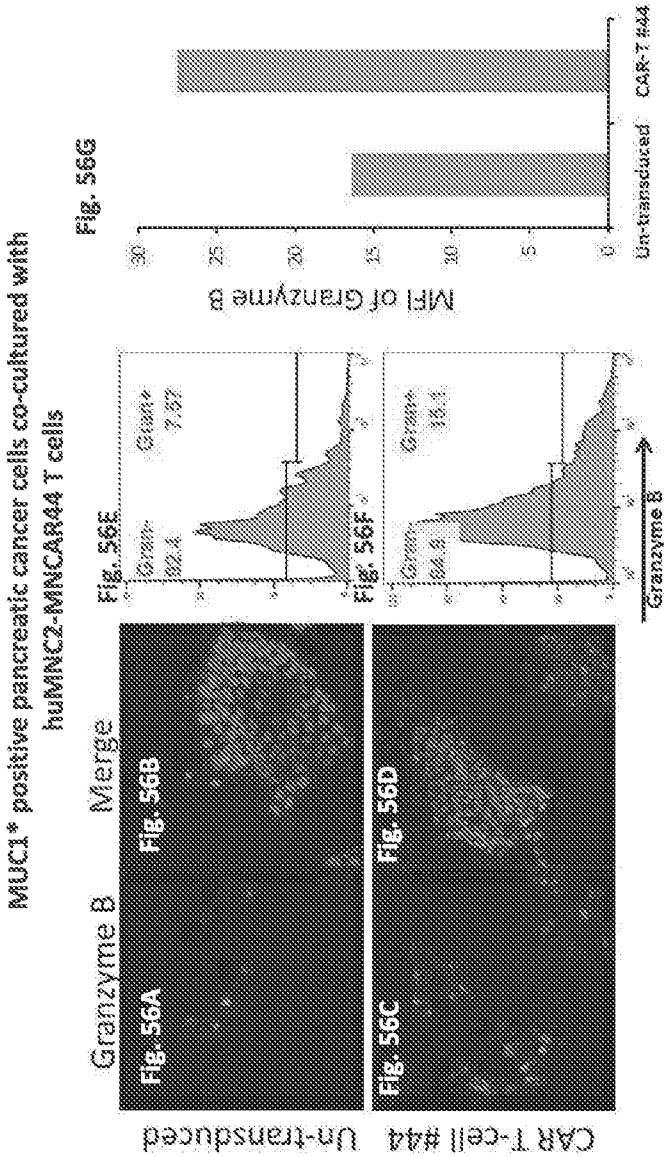


Figure 55A-55H



CAPAN-2

Fig. 56H

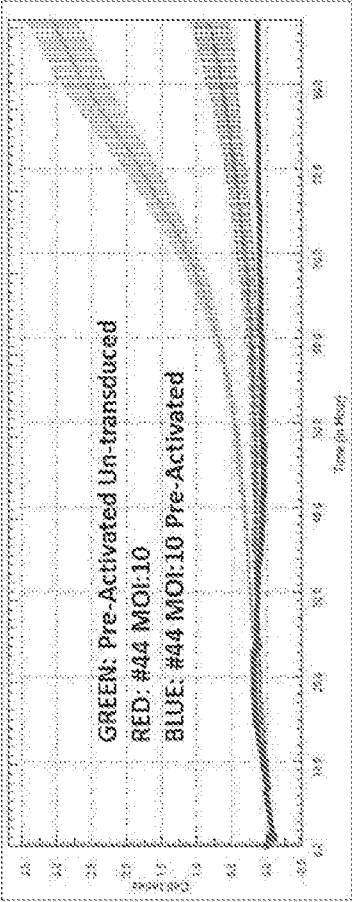


Figure 56A-56H



Fig. 57B

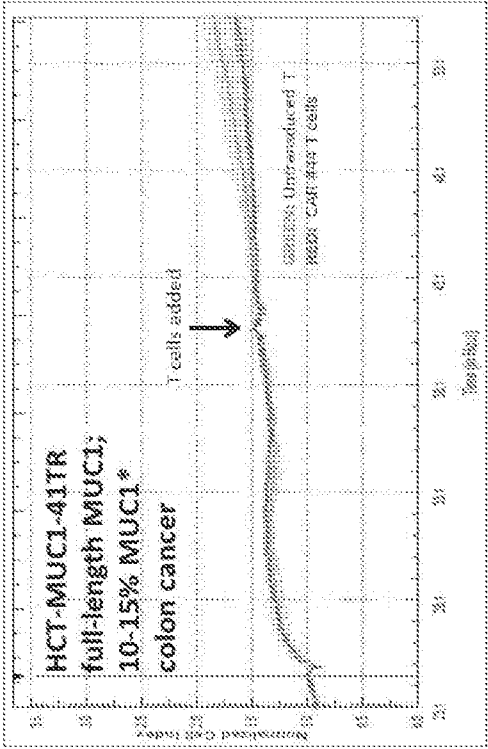


Fig. 57A

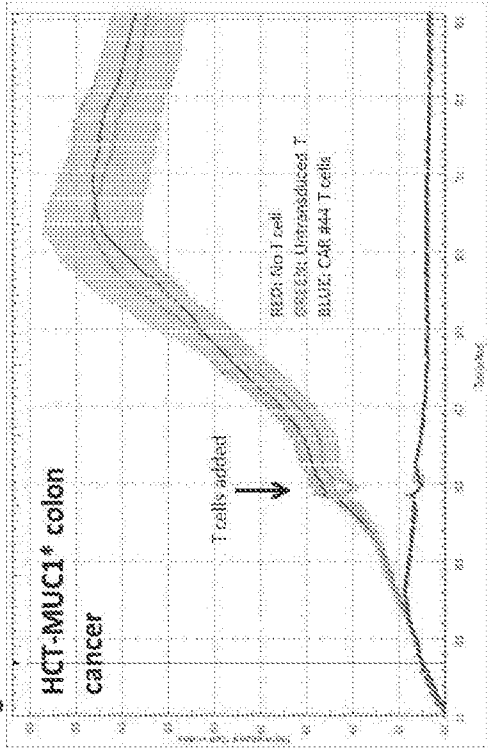


Fig. 57C

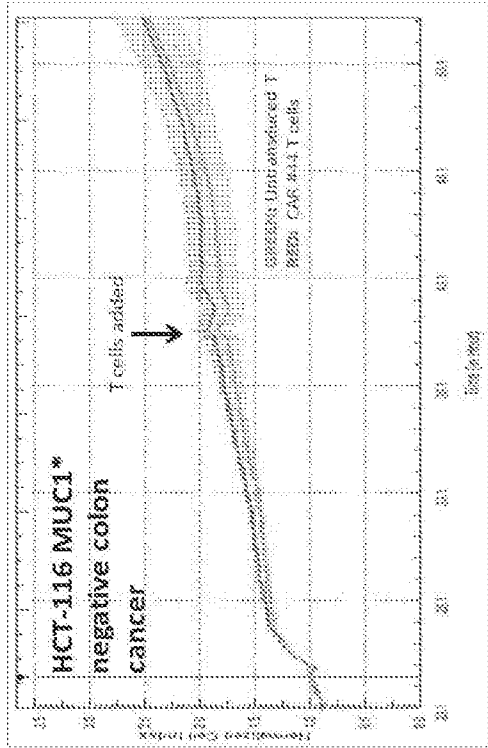


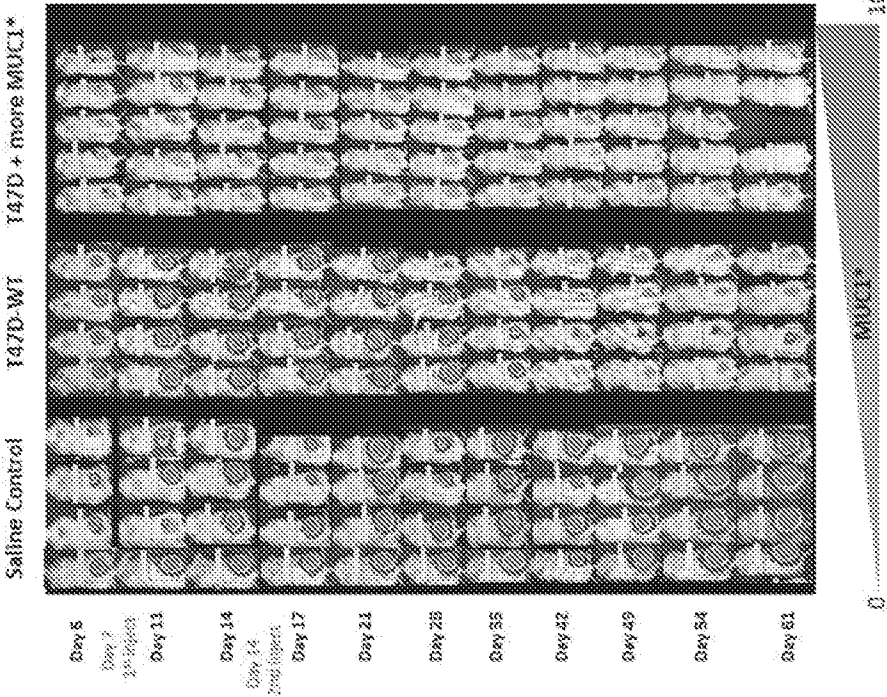
Figure 57A-57C



huMNC2-CAR44 T kill human breast tumors (n > 300)

Killing increases as MUC1\* expression increases; MUC1\* expression increases with tumor stage & acquired resistance to chemo drugs

Fig. 59A                      Fig. 59B                      Fig. 59C



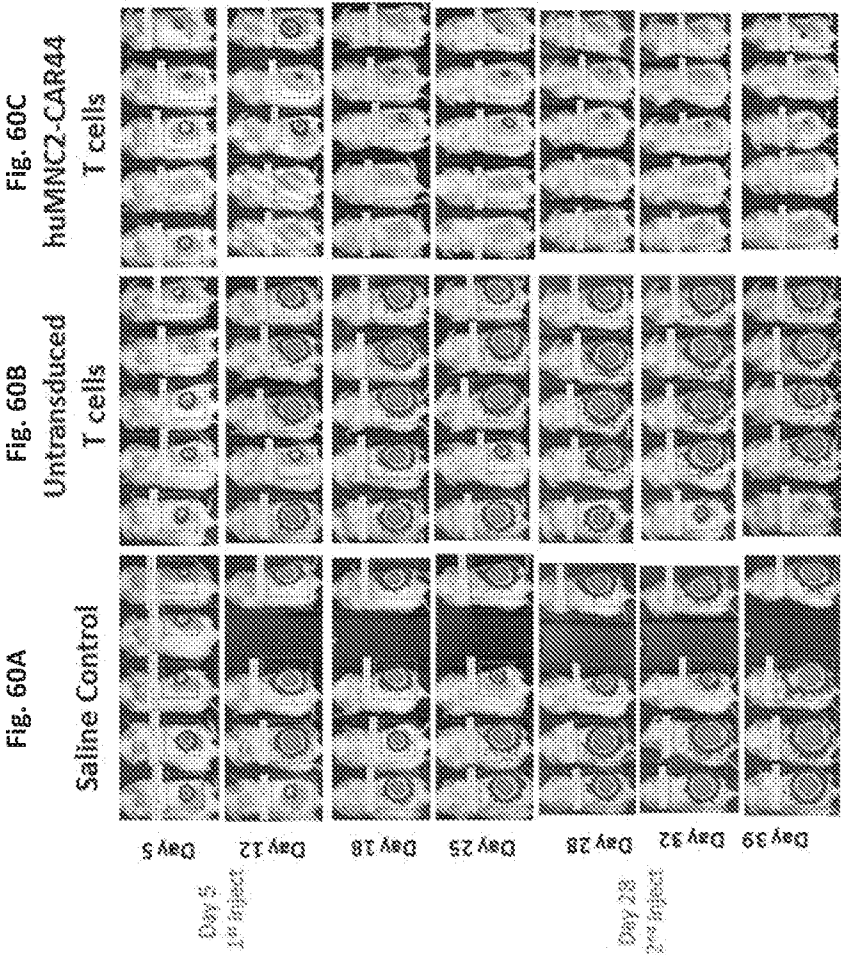
T47D-wt is a naturally occurring metastatic breast cancer cell line

T47D + more MUC1\* is that same cell line but with 95% of the tumor cells engineered to express more MUC1\*

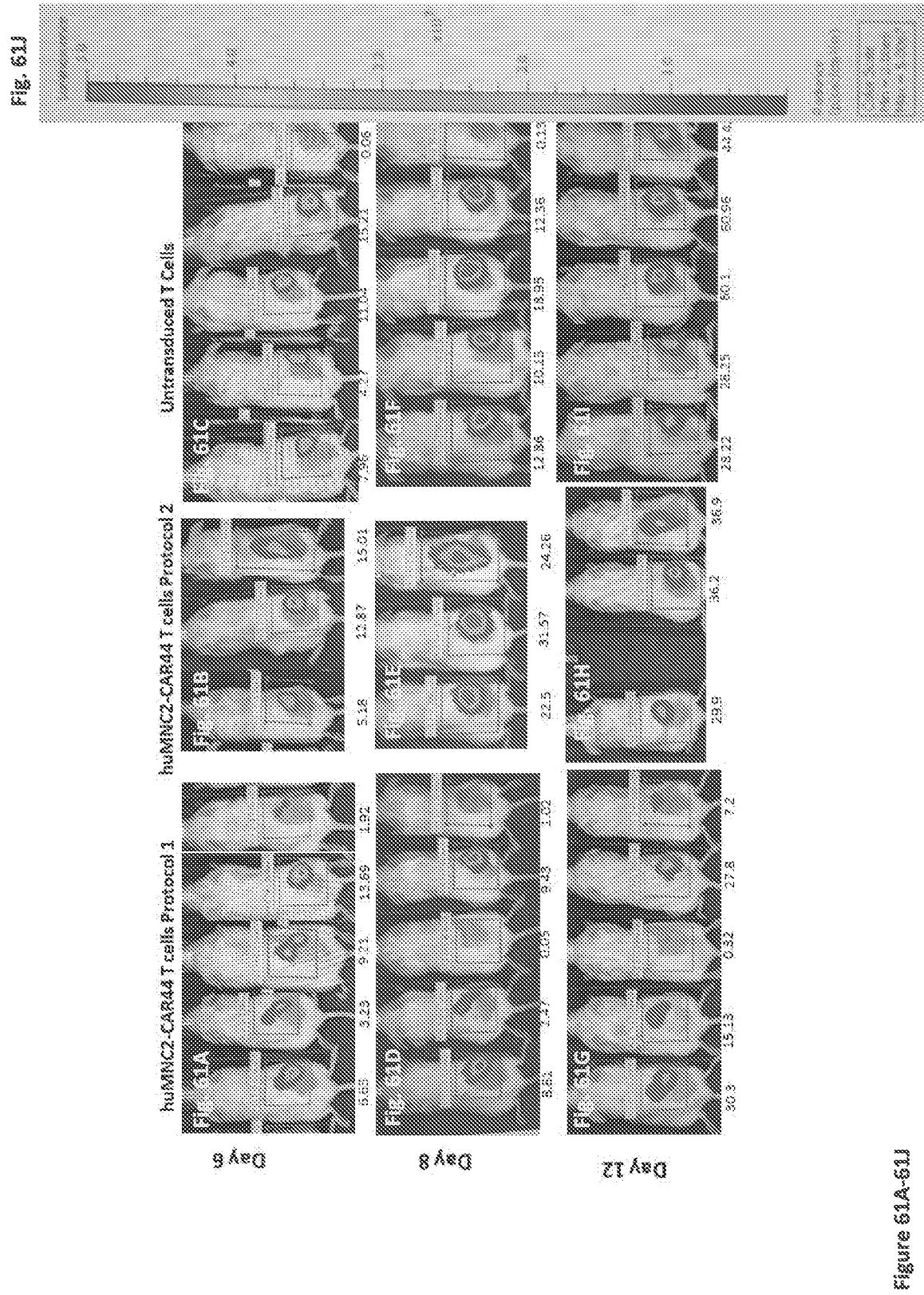
Figures 59A-59C

**huMNC2-CAR44 T cells kill breast tumors even if only a low percentage of tumor expresses high amounts of MUC1\***

T47D human metastatic breast tumors where 30% of the tumor cells were engineered to express more MUC1\* mimics mid stage tumors



Figures 60A-60C



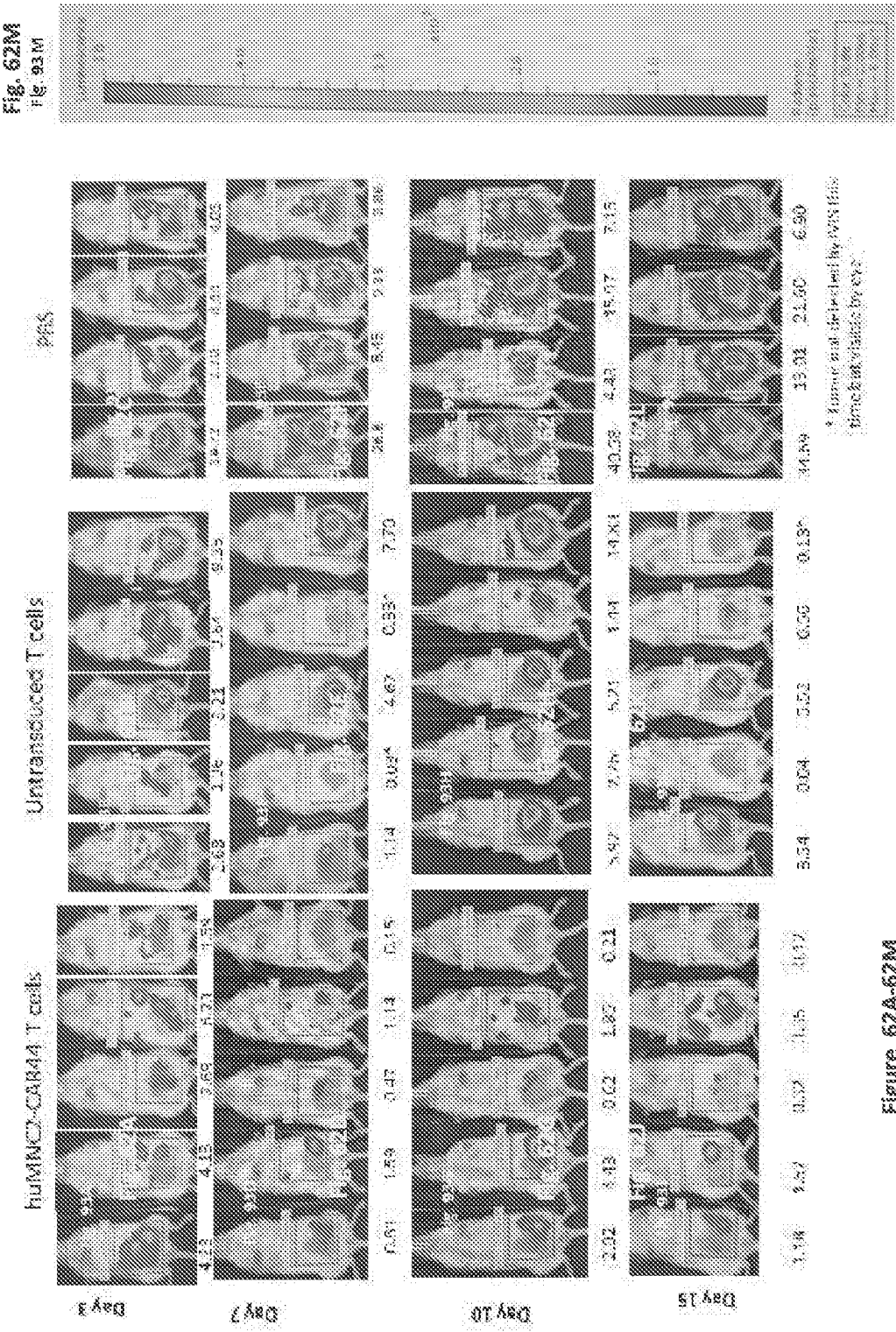


Figure 62A-62M

ELISA assay tests ability of monoclonal antibodies to bind to various peptides whose sequences are derived from the PSMGFR peptide

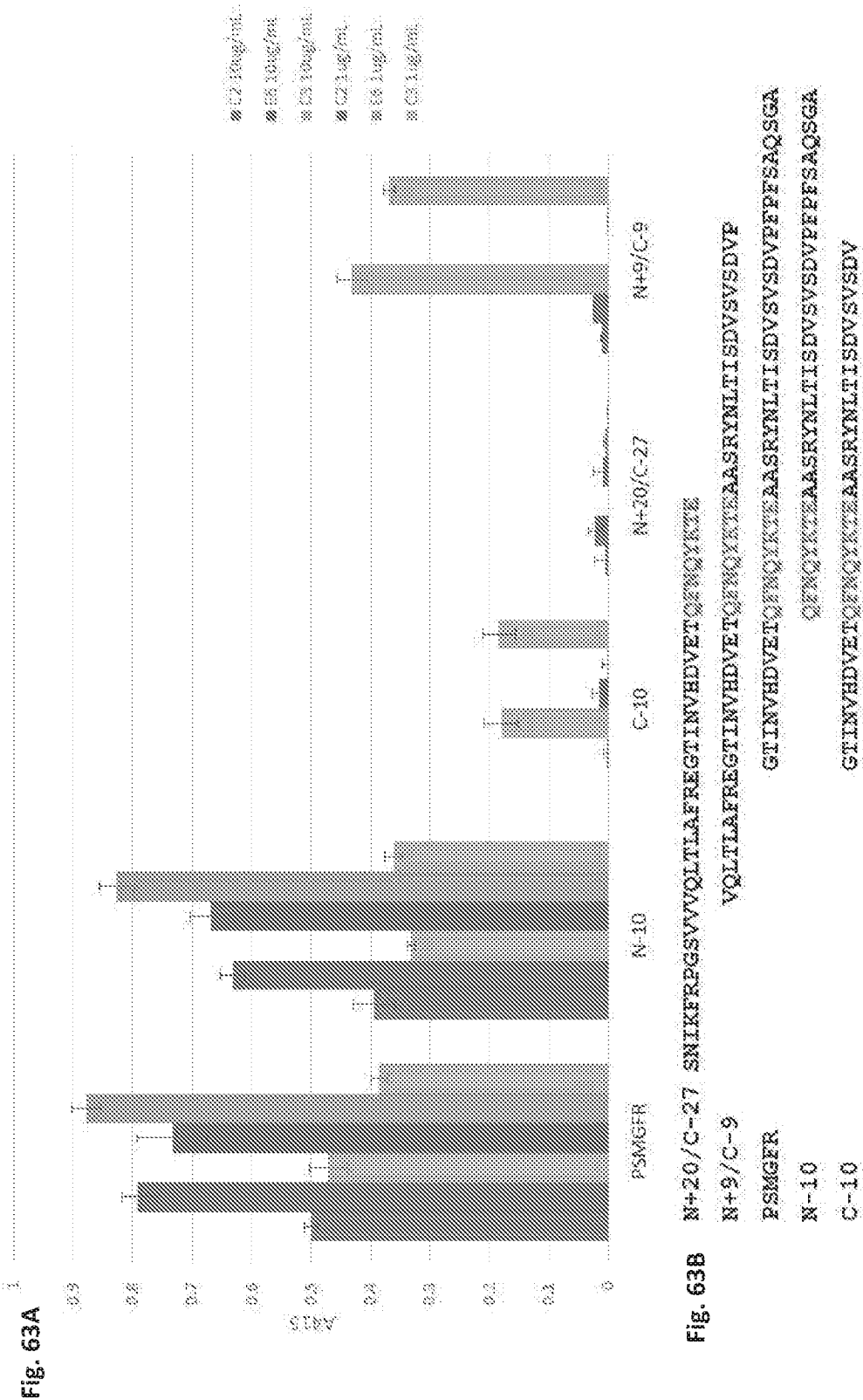


Figure 63A-63B

Animals immunized with PSMGFR peptide; 1<sup>st</sup> selection: bind to PSMGFR; color indicates deductive cognate epitope

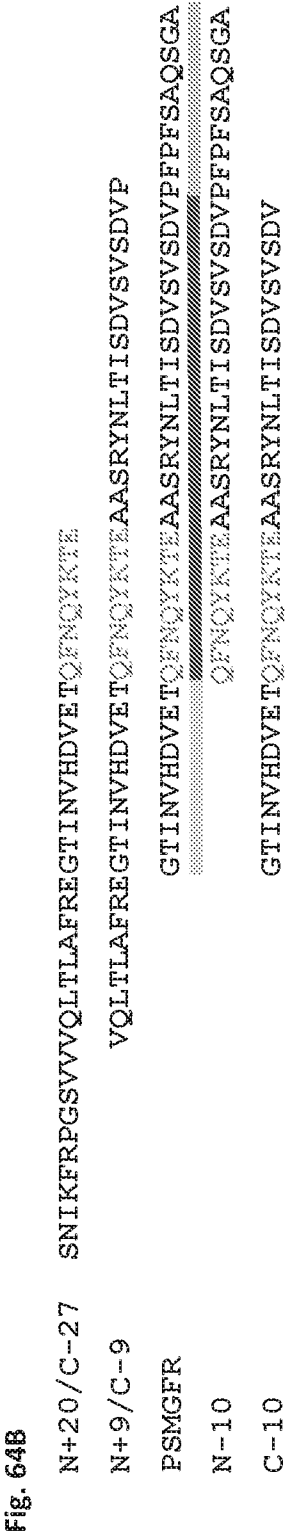
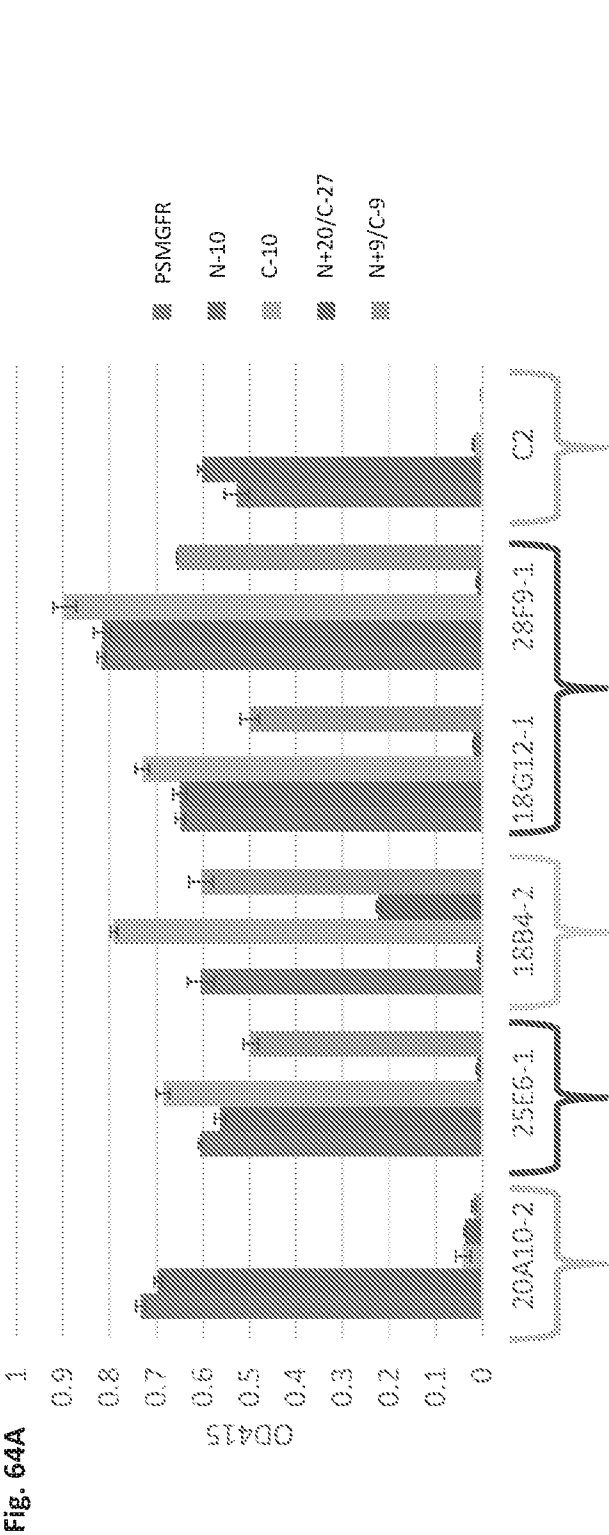


Figure 64A-64B



Animals immunized with N+20/C-27 peptide; 1st selection: binds to N+20/C-27; color indicates deductive cognate epitope

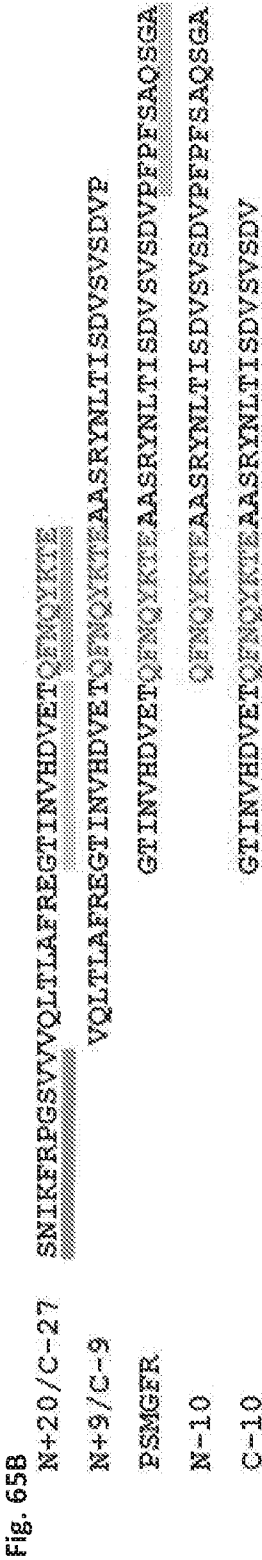
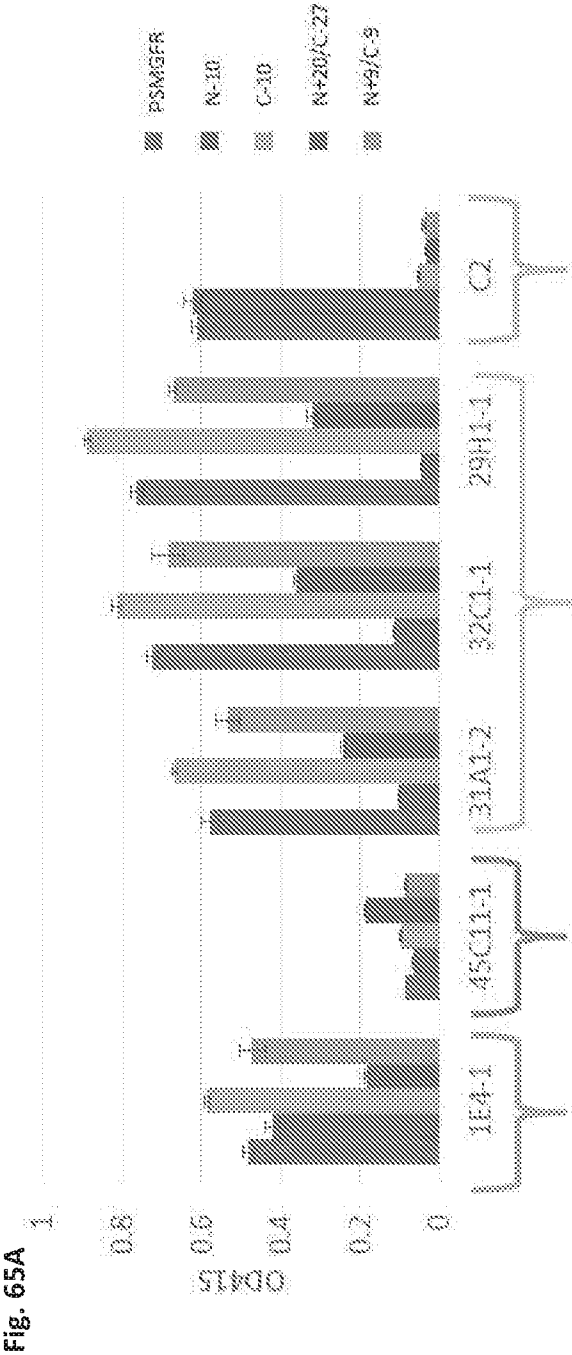


Figure 65A-65B

Animals immunized with 9+C-9 peptide; 1st selection: bind to 9+C-9; color indicates deductive cognate epitope

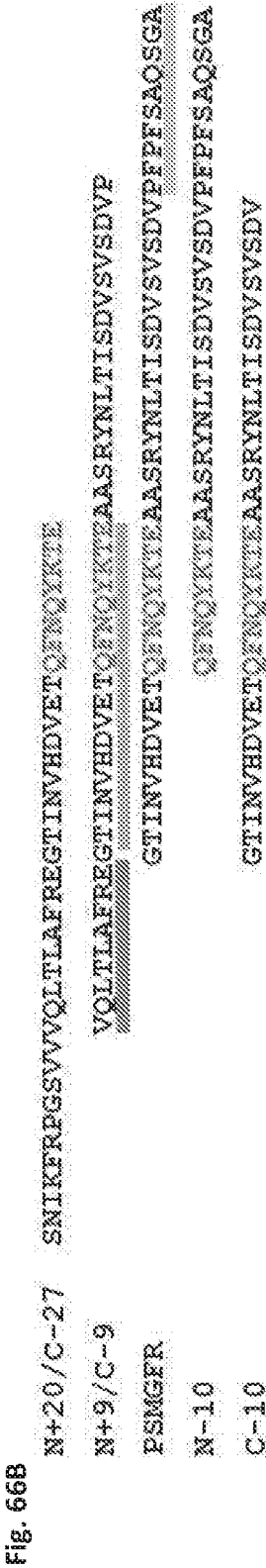
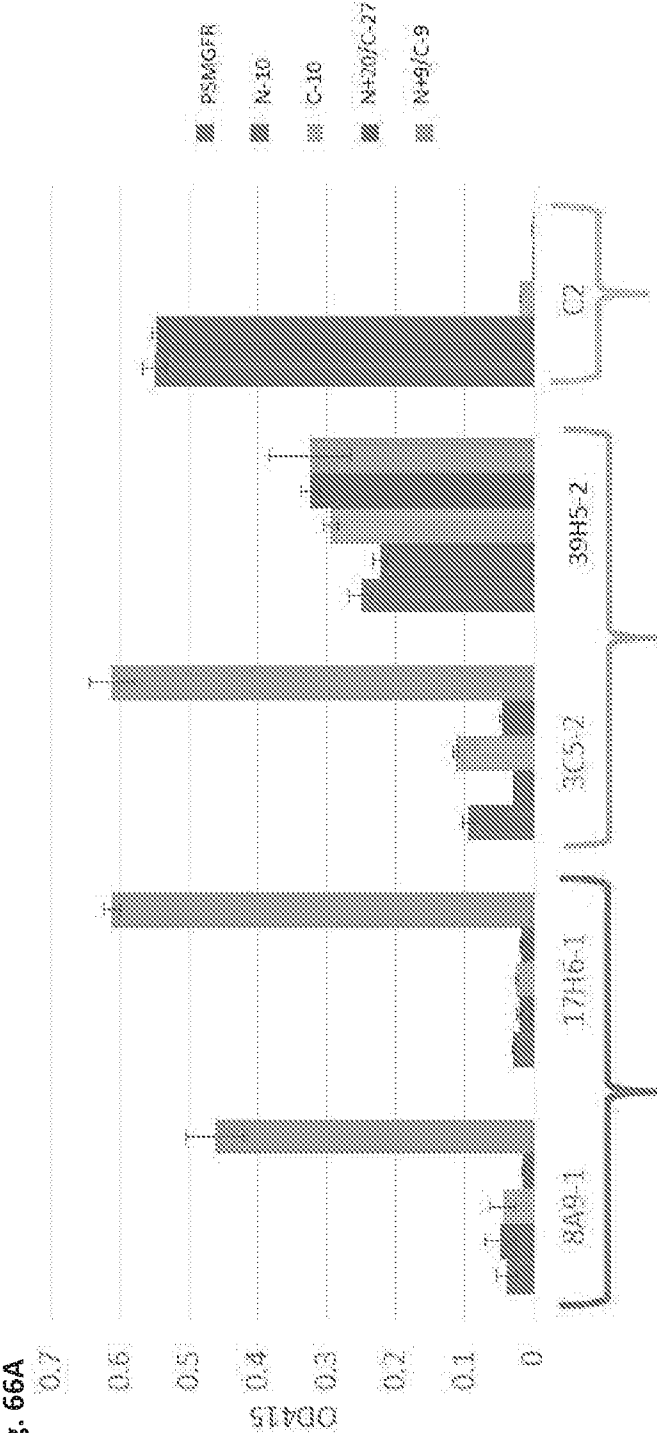


Figure 66A-66B

ELISA: PSMGFR antibodies – refined epitope mapping (all antibodies @10ug/mL)

Fig. 67A

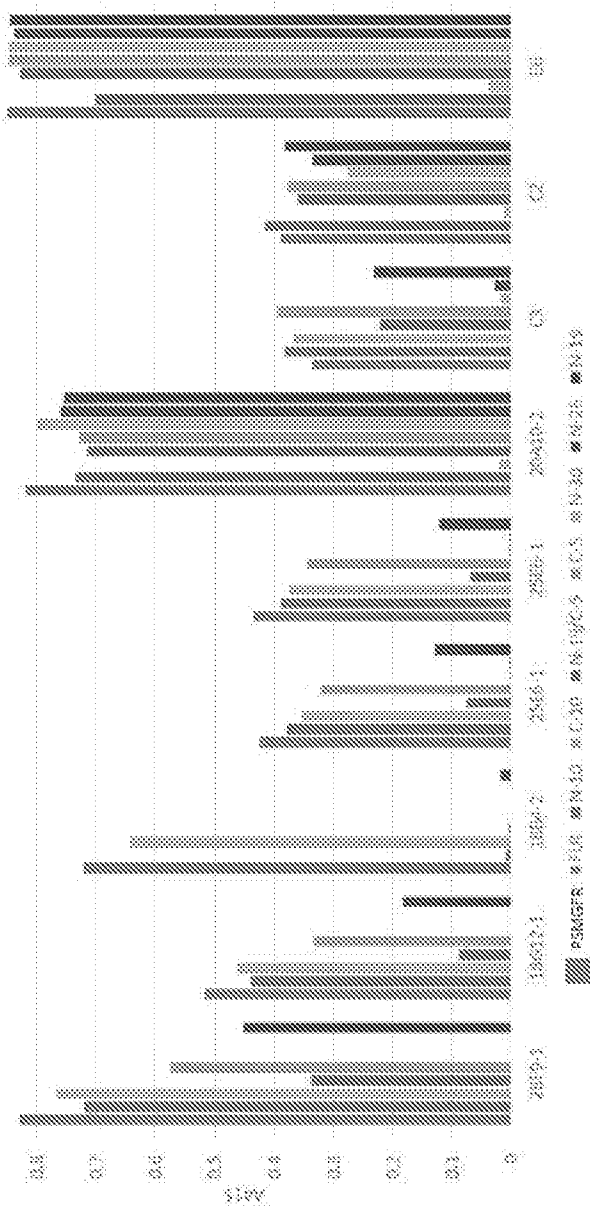


Fig. 67B

SNIKRPGSVVQLTLAFREGTINVHVDVETQFNQYKTEAASRYNLTISDVSVSDVPEPFPSAQSGA

Fig. 67C

N-19	ASRYNLTISDVSVSDVPEPFPSAQSGA	Refined epitope	ASRYNLT
N-26	ISDVSVSDVPEPFPSAQSGA	2679-1	ASRYNLT
N-30	SVSDVPEPFPSAQSGA	18612-1	GTINVHVDVET
N-10/C-5	QFNQYKTEAASRYNLTISDVSVSDVPEPFPS	1834-2	ASRYNLT
N-19/C-5	ASRYNLTISDVSVSDVPEPFPS	2526-1	ASRYNLT
PSMGFR	GTINVHVDVETQFNQYKTEAASRYNLTISDVSVSDVPEPFPSAQSGA	20410-2	PSMGFR
N-10	QFNQYKTEAASRYNLTISDVSVSDVPEPFPSAQSGA	C3	ASRYNLT
C-10	GTINVHVDVETQFNQYKTEAASRYNLTISDVSVSDV	C1	PSMGFR
		C0	PSMGFR

Figure 67A-67D

ELISA: N+20/C-27 antibodies – refined epitope mapping (all antibodies @10ug/mL)

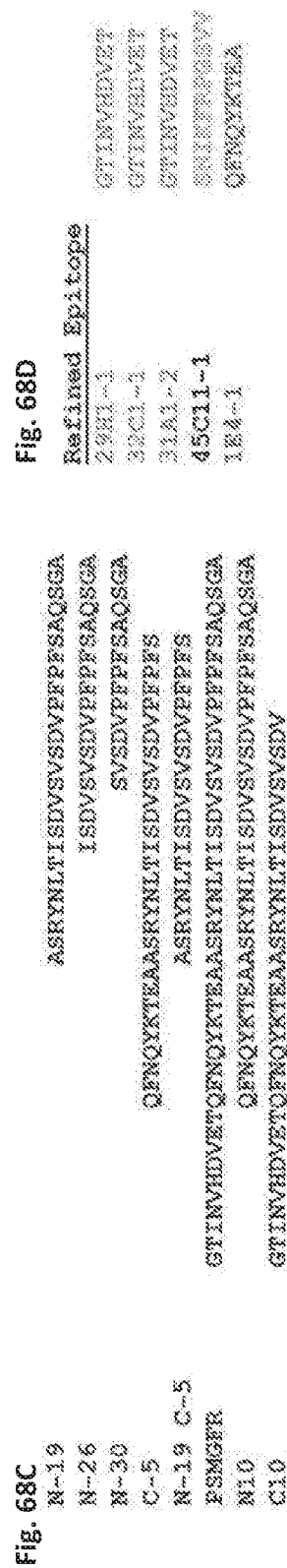
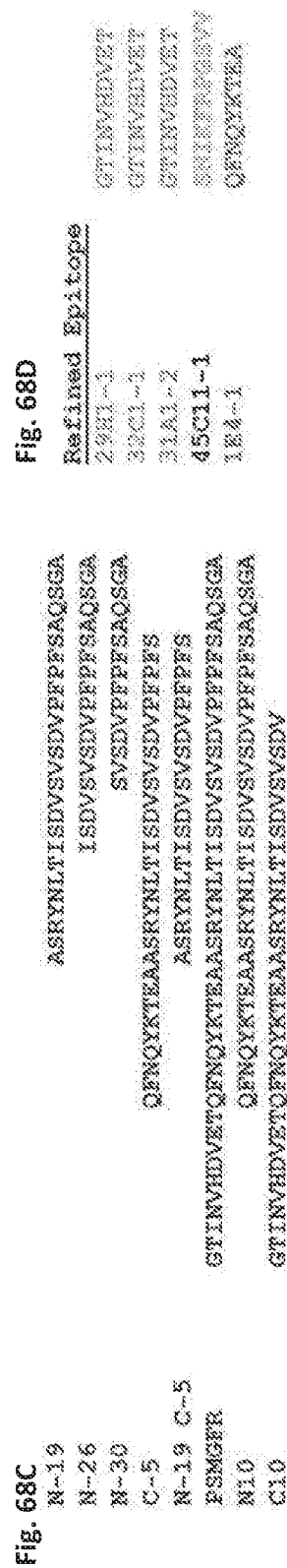
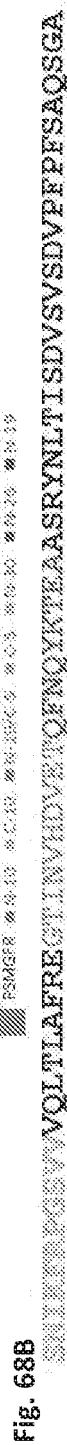
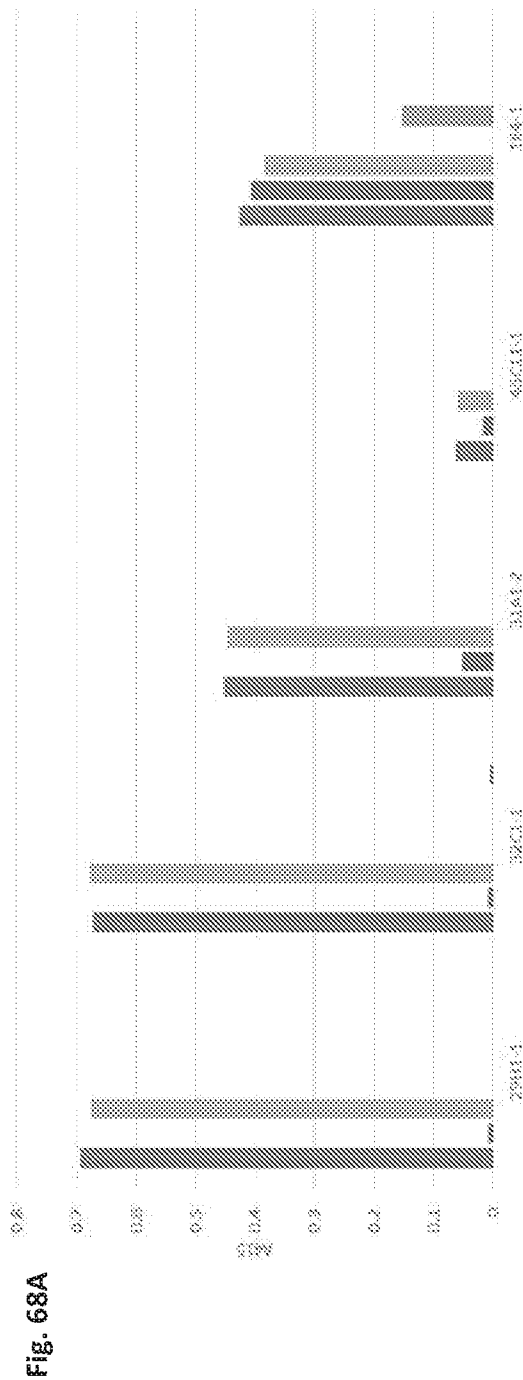


Figure 68A-68D

ELISA: N+9/C-9 antibodies – refined epitope mapping (all antibodies @10ug/mL)

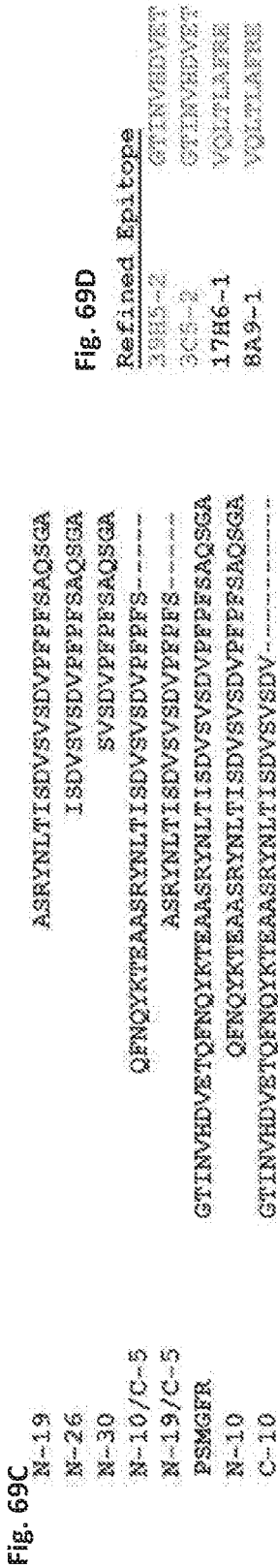
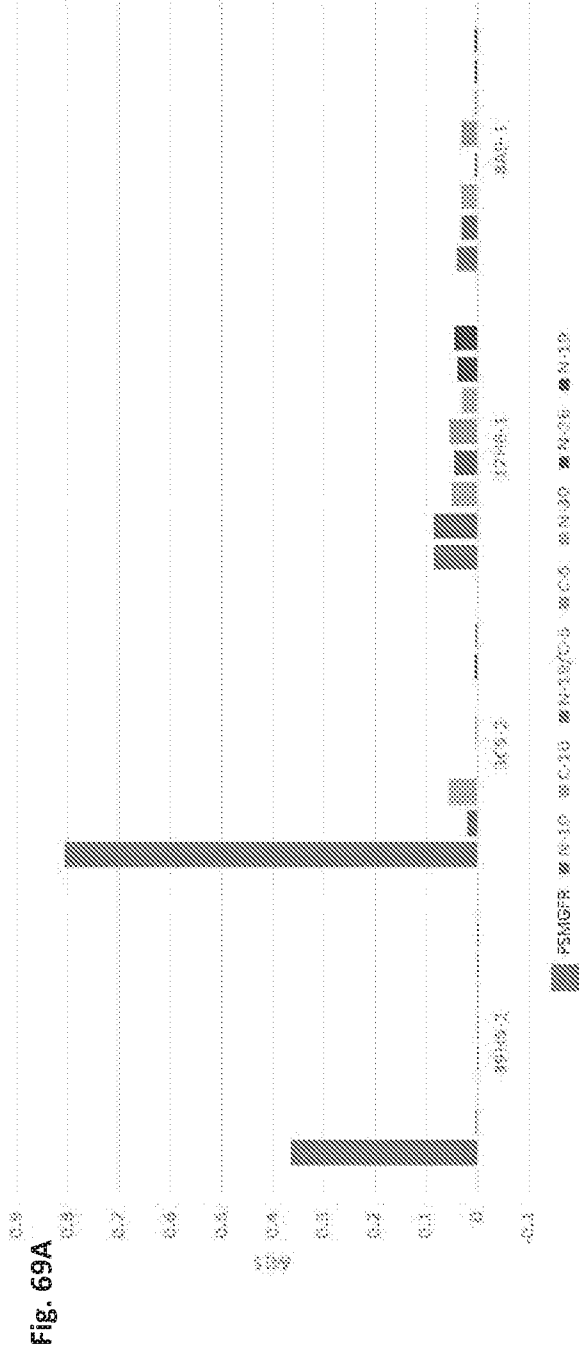


Figure 69A-69D

ELISA displacement assay tests ability of antibodies of the invention to displace NME7<sub>AB</sub> from binding to PSMGFR peptide.

Figure 70A

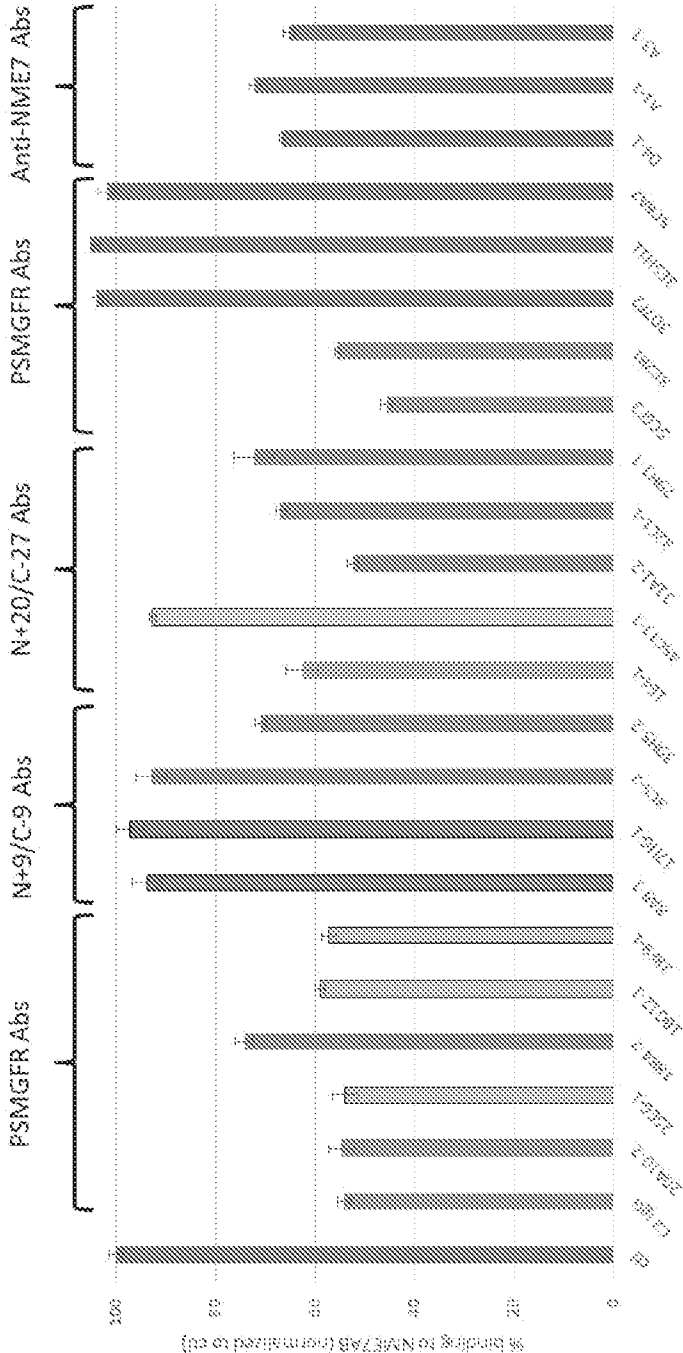


Figure 70B

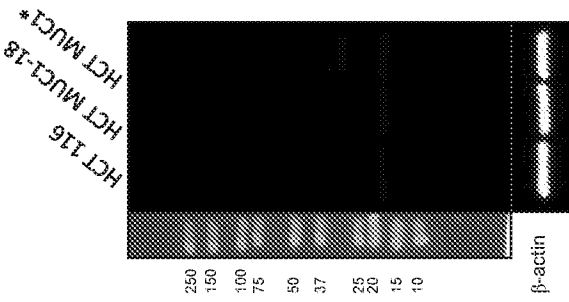
SNIKFRPGSVVQLTLAFREGTINVDVETQFNQYKTEAASRYNLTISDVSVSDVPPPPSAQSGA

Figure 70A-70B

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Fig. 71A

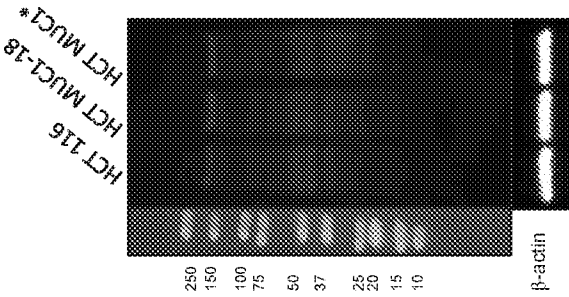
MN-C2



Stock Concentration: 1.85 mg/mL  
Dilution Tested: 1:500  
Exposure: 5 min  
Reagent: ECL

Fig. 71B

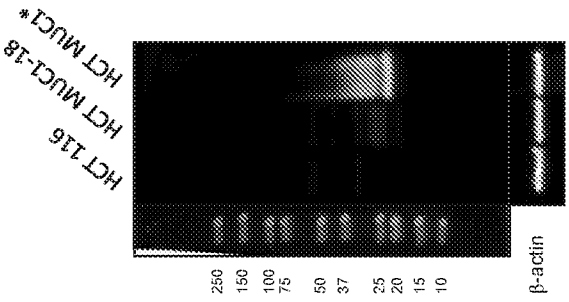
MN-E6



Stock Concentration: 1.859mg/mL  
Dilution Tested: 1:500  
Exposure: 5 min  
Reagent: ECL

Fig. 71C

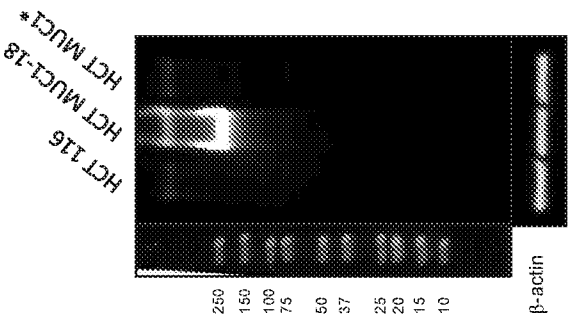
SDIX



Stock Concentration: 1mg/mL  
Dilution : 1:1000  
Exposure: 2 min  
Reagent: ECL

Fig. 71D

VU4H5



Stock Concentration: 200 µg/mL  
Dilution : 1:200  
Exposure: 5 min  
Reagent: ECL

Figure 71A-71D

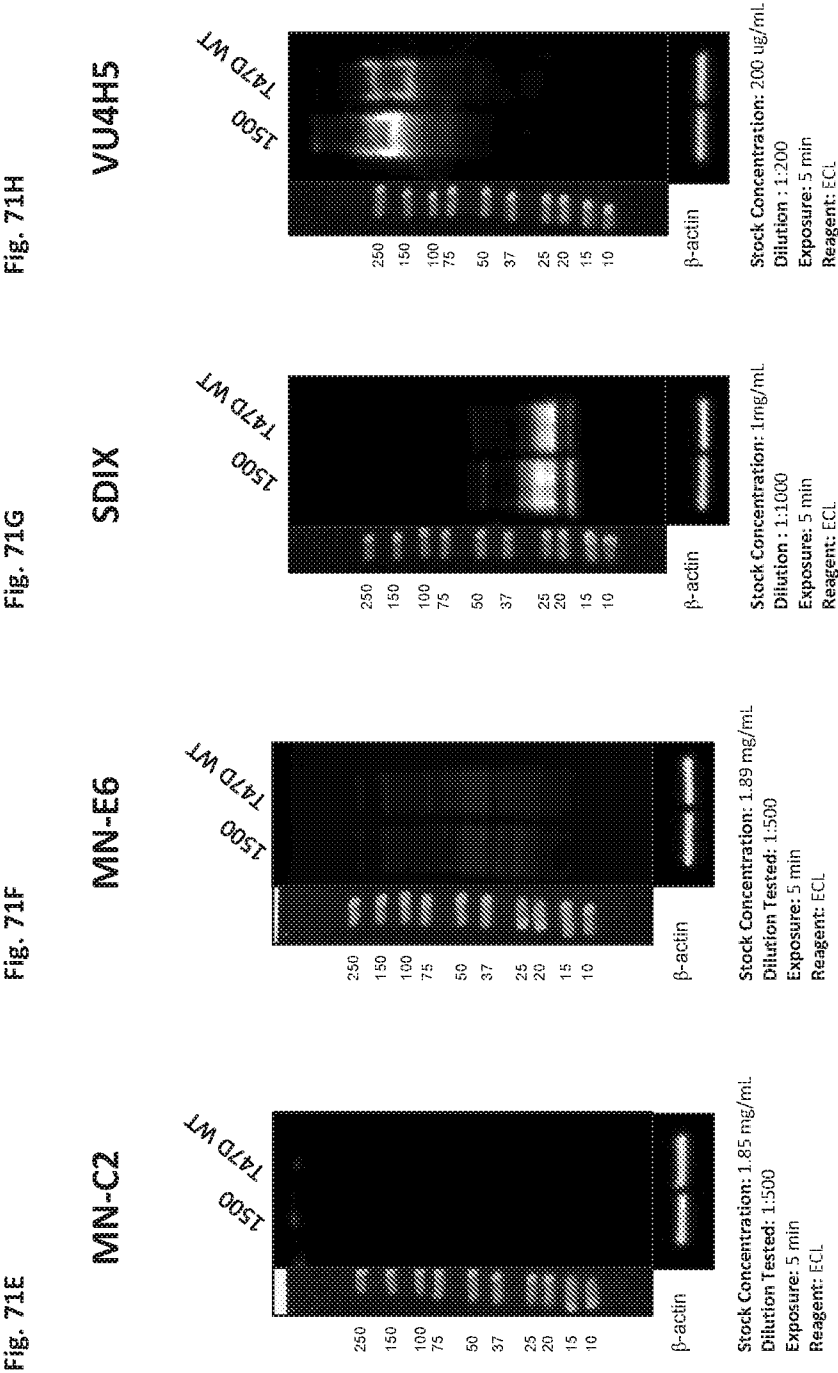


Figure 71E-71H



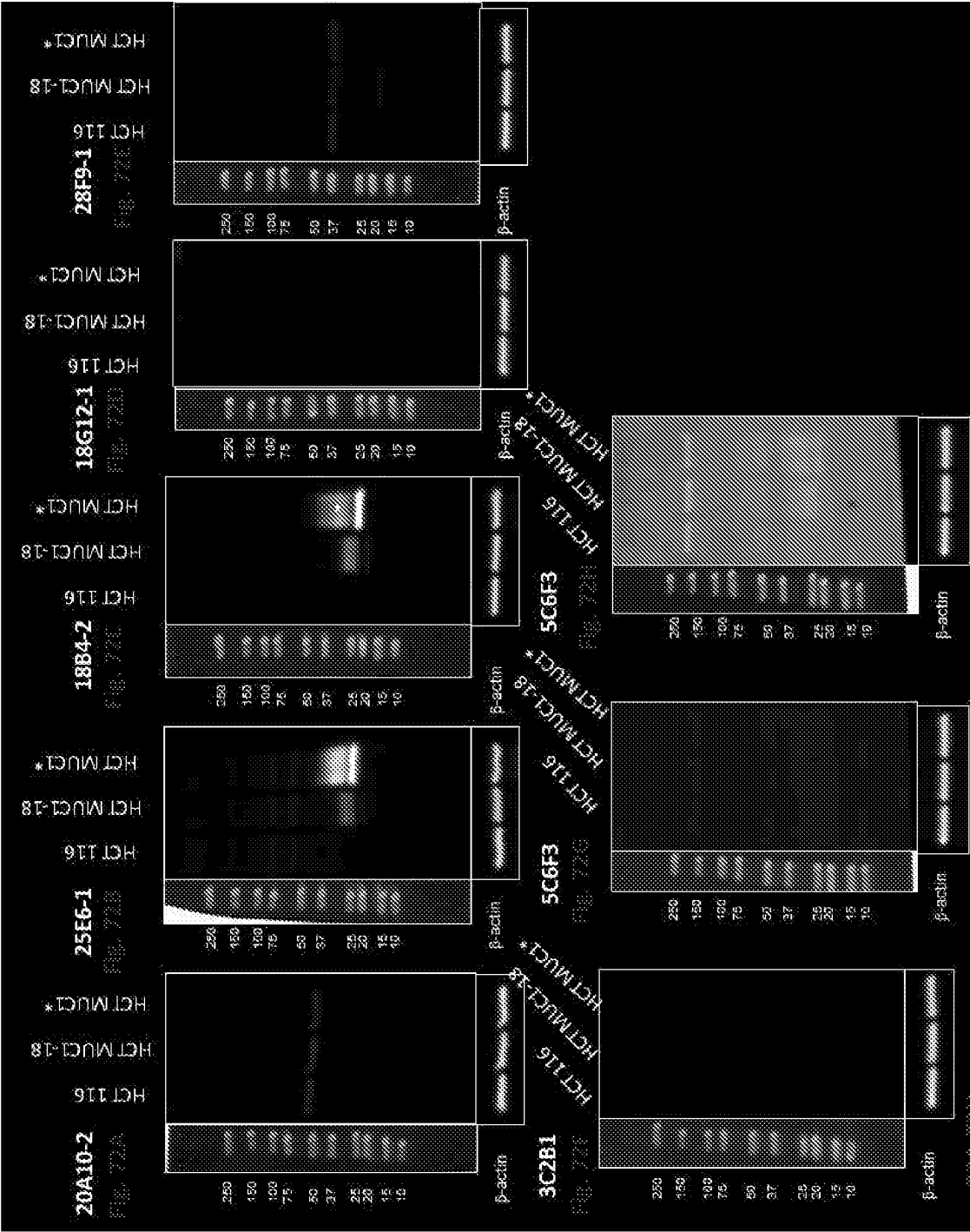
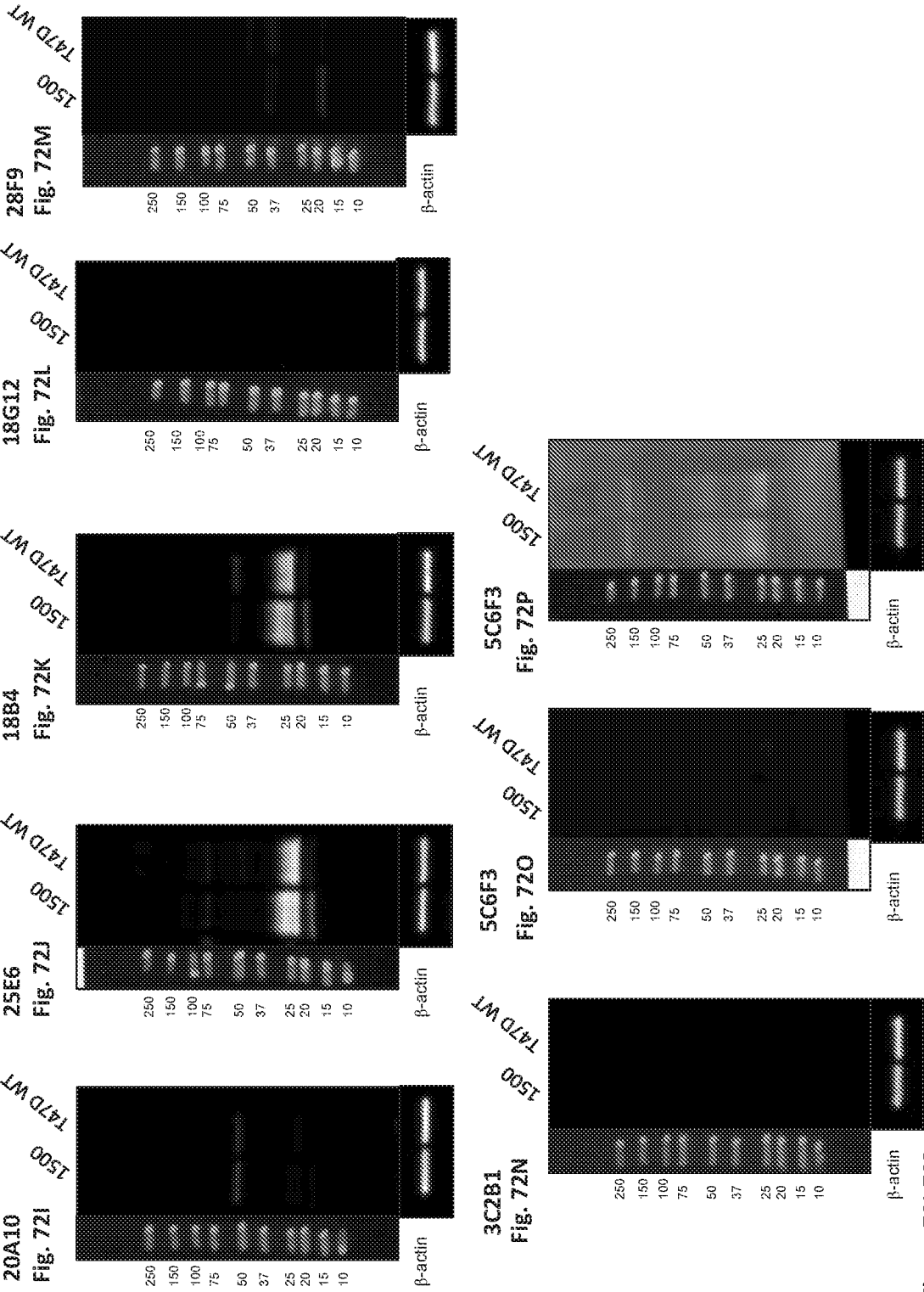


Figure 72A-72H



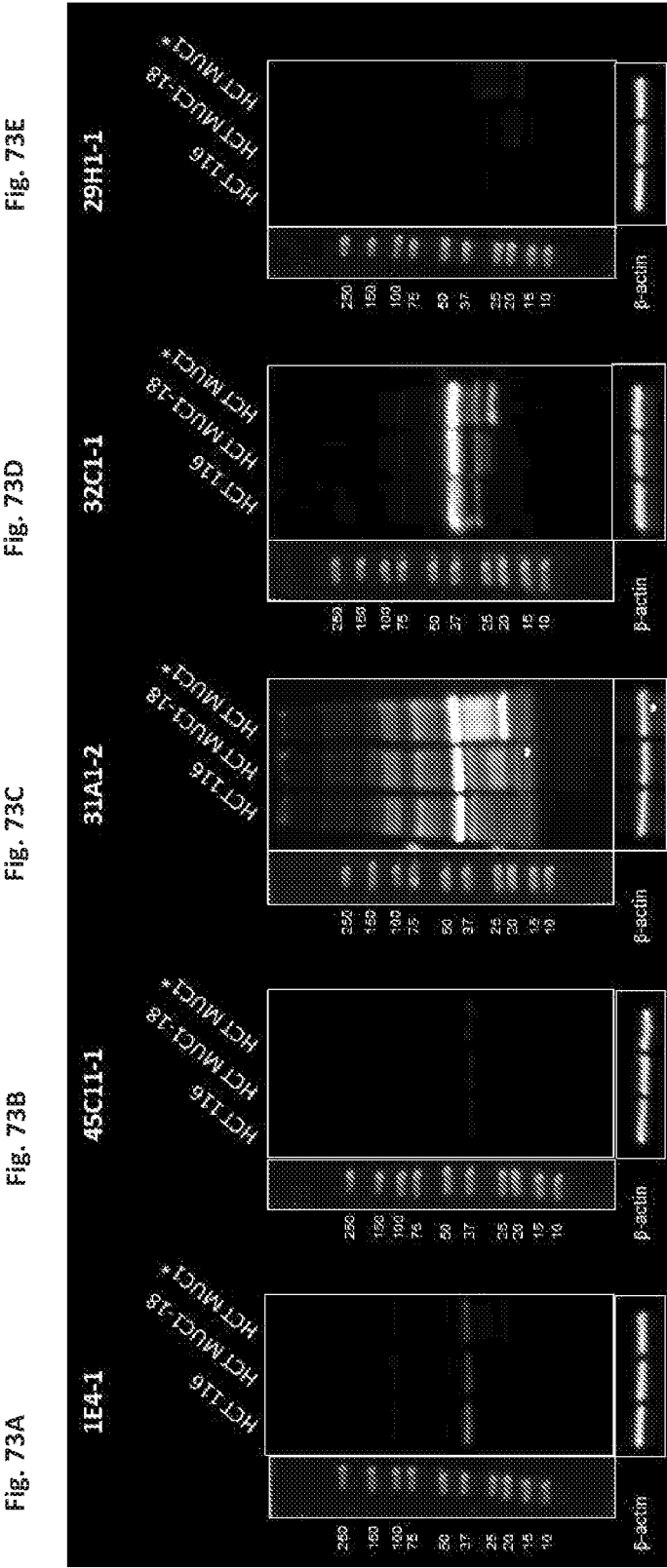


Figure 73A-73E

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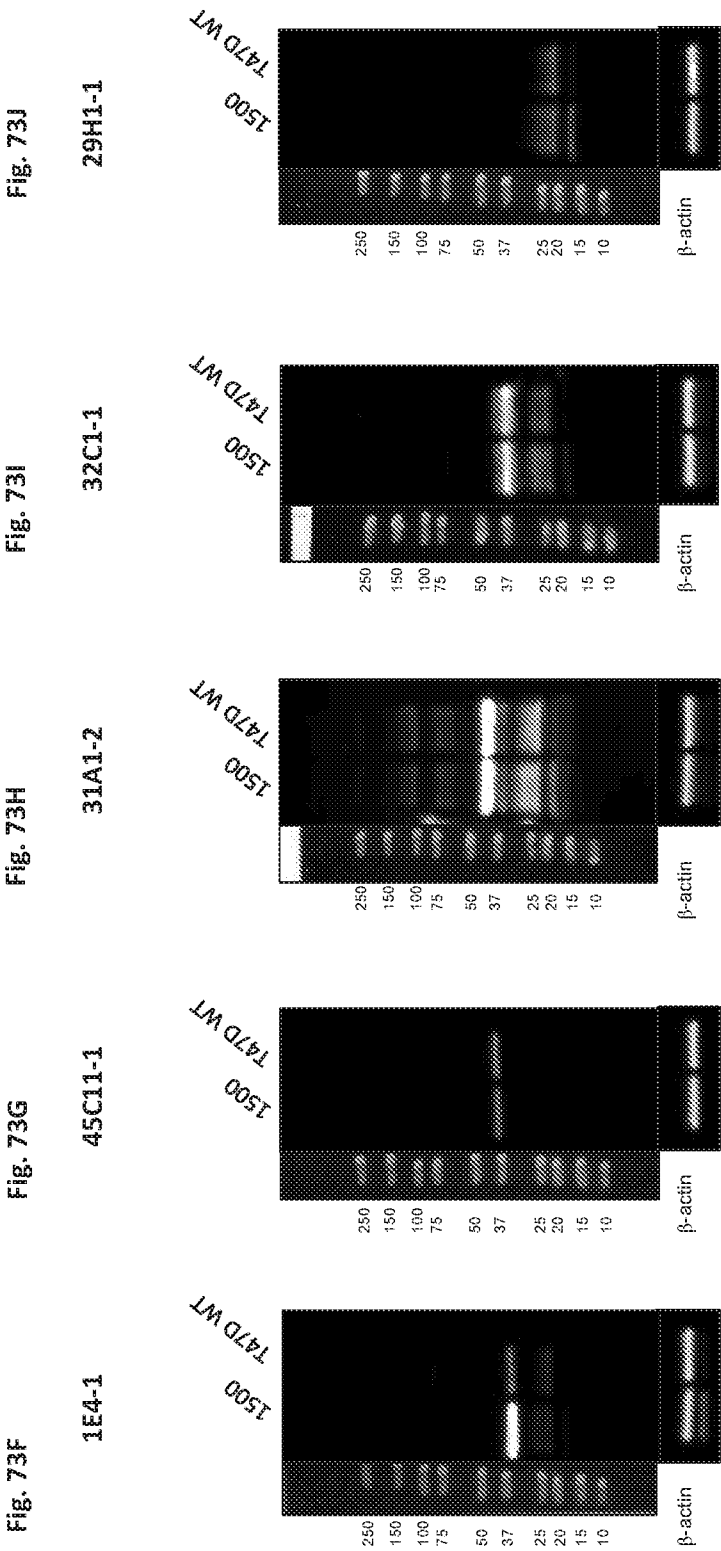


Figure 73F-73J

Fig. 74A

8A9-1

HCT 116  
HCT MUC1-18  
\*HCT MUC1\*

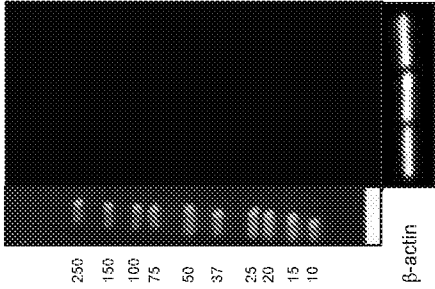


Fig. 74B

17H6-1

HCT 116  
HCT MUC1-18  
\*HCT MUC1\*

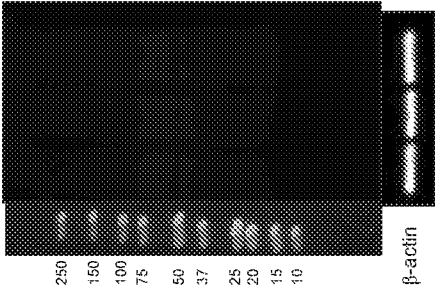


Fig. 74C

3C5-2

HCT 116  
HCT MUC1-18  
\*HCT MUC1\*

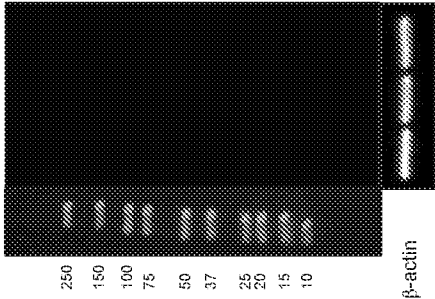


Fig. 74D

39H5-2

HCT 116  
HCT MUC1-18  
\*HCT MUC1\*

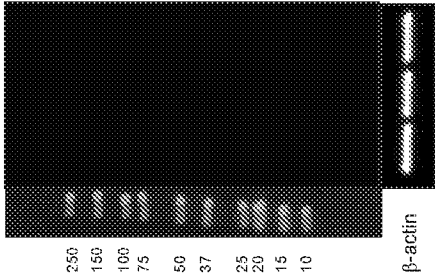


Figure 74A-74D

Fig. 74E

8A9-1

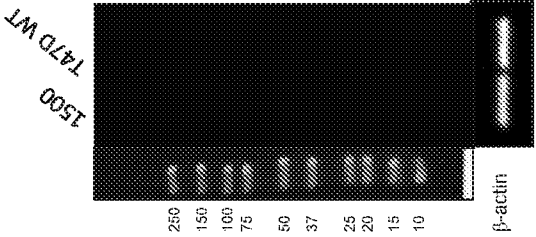


Fig. 74F

17H6-1

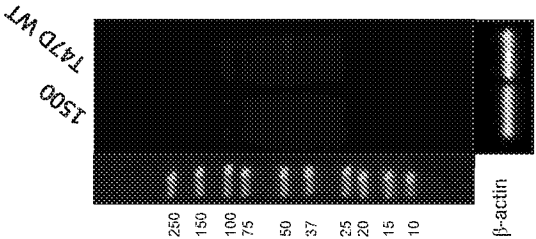


Fig. 74G

3C5-2

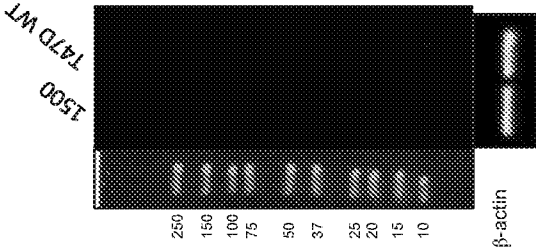


Fig. 74H

39H5-2

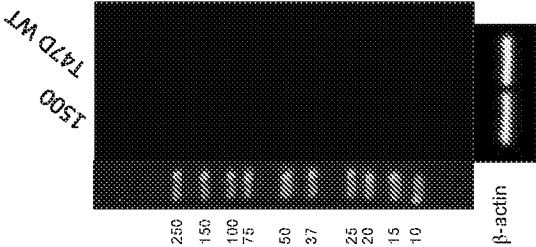


Figure 74E-74H

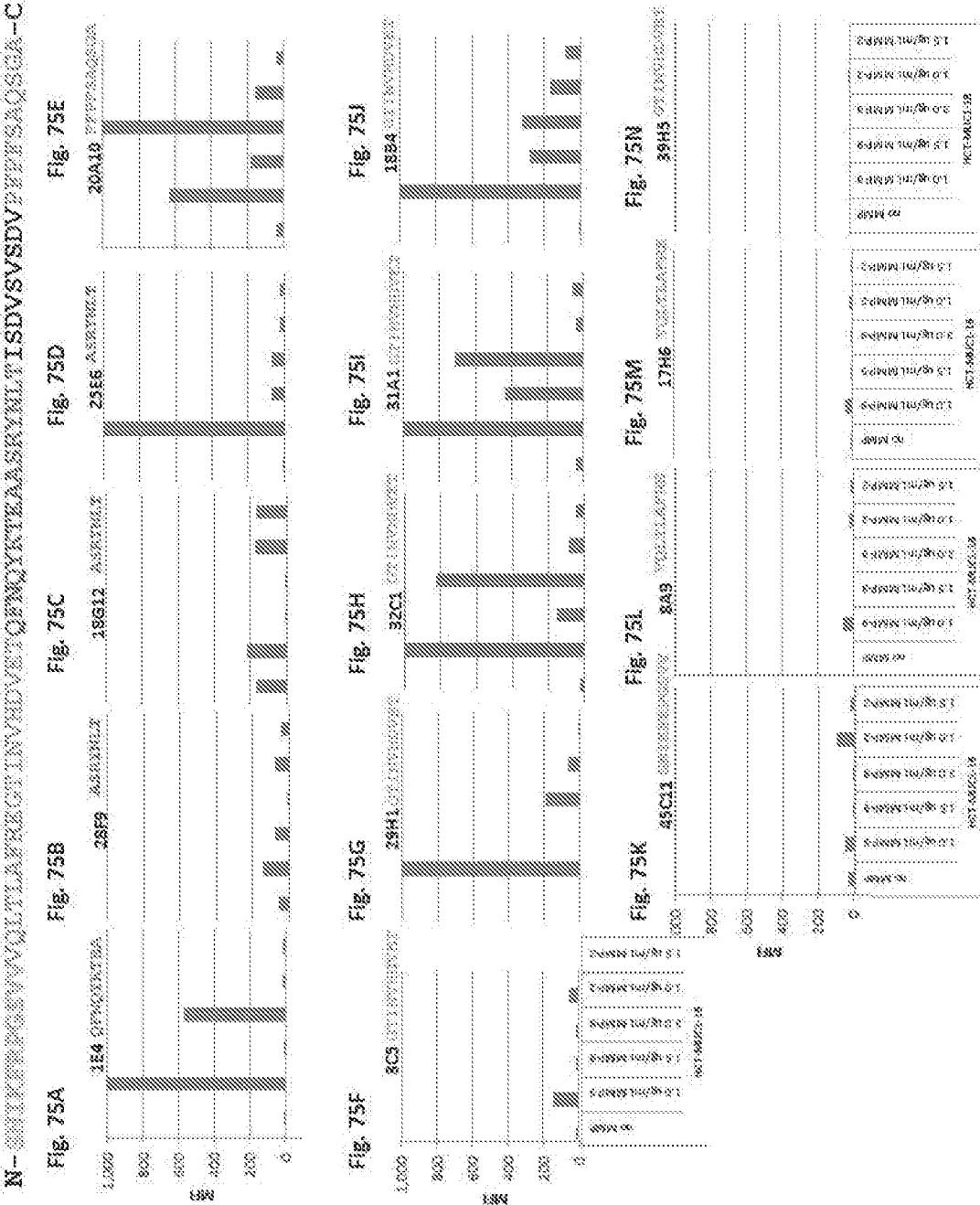


Figure 75A-75N

N-SNKKRPGSW VQLTLAFRE GTINVDVET QFNQYKTEA ASRYNLT ISDVSVSDV PPFESAQSGA -C

Fig. 750

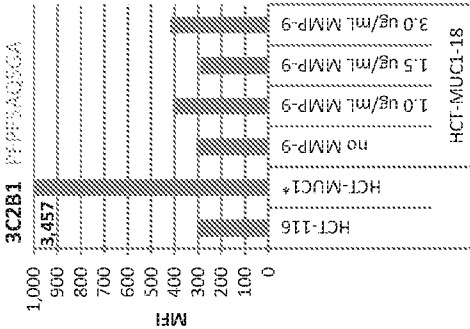


Fig. 75P

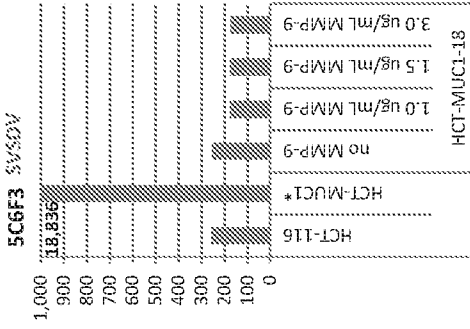


Figure 750-75P



FACS: Reference antibodies per cell line

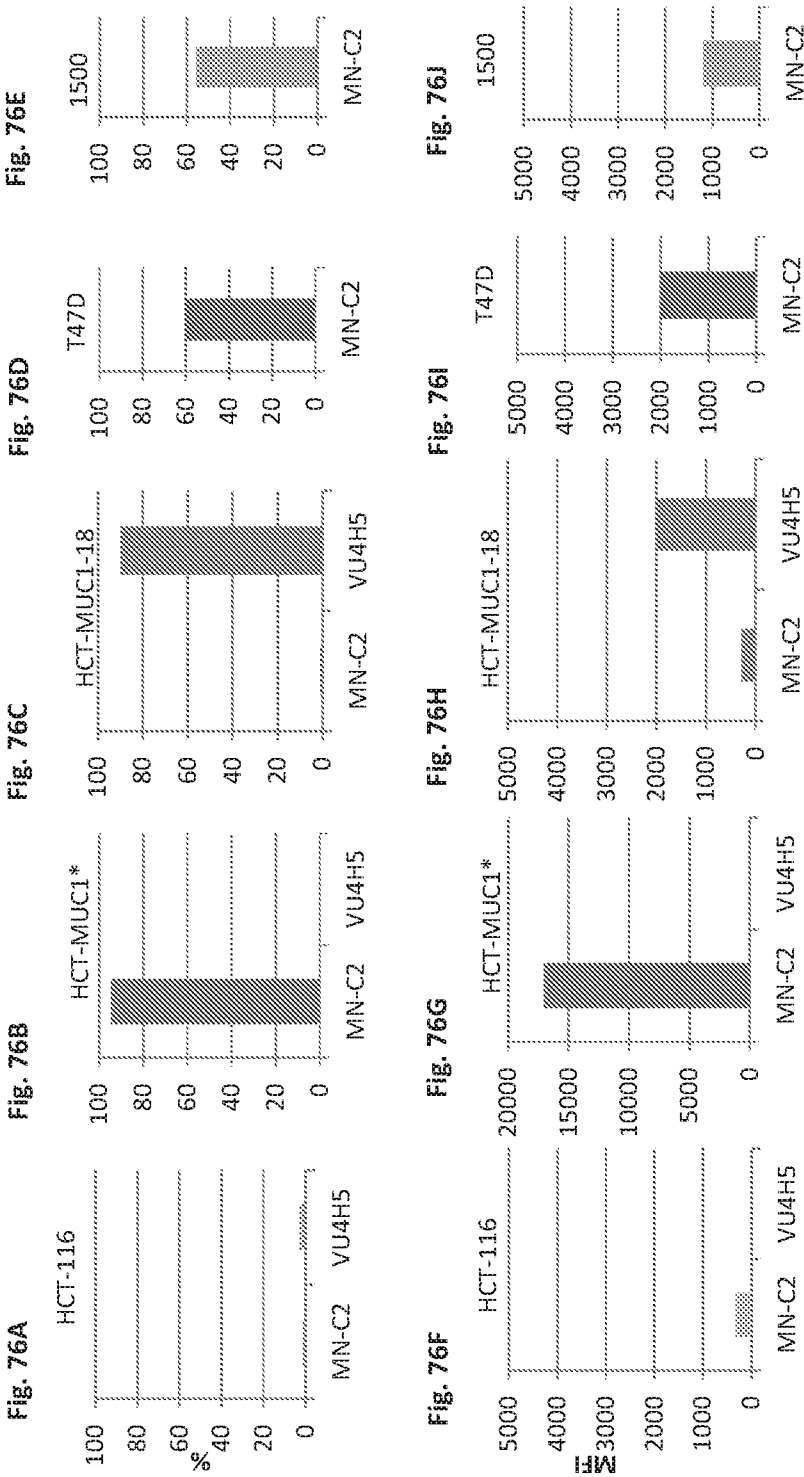


Figure 76A-76J

FACS: Reference antibodies per cell line

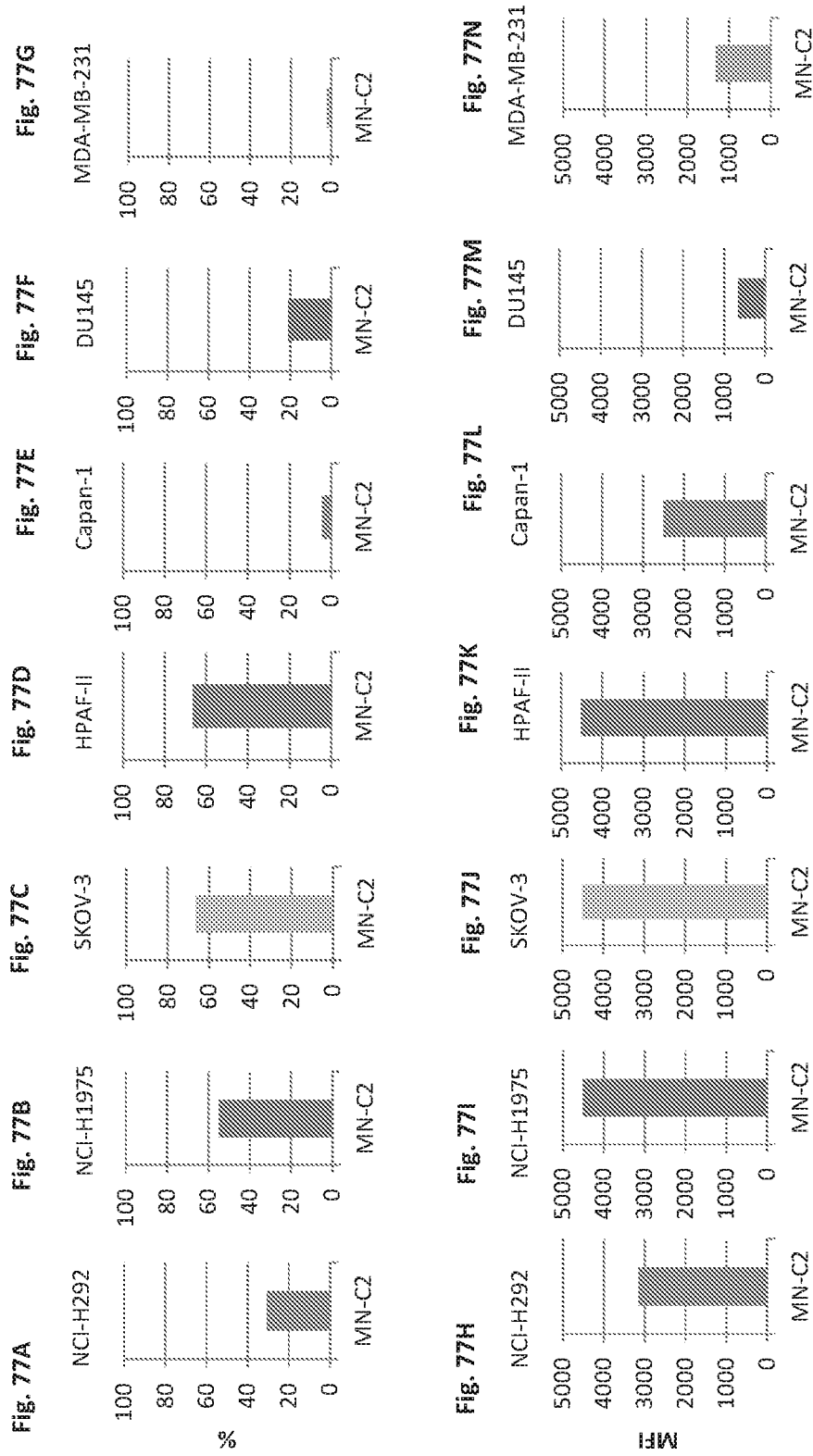


Figure 77A-77N

Fig. 78A

PSMGR	SHIKTRGGSVAVQLTLAFREGTINVDVETQFNQKTEAASRYMLTISDVSVSDVPFPFSAQSGA	
N+20/C-27	GTINVDVETQFNQKTEAASRYMLTISDVSVSDVPFPFSAQSGA	
N+9/C-9	SHIKTRGGSVAVQLTLAFREGTINVDVETQFNQKTEAASRYMLTISDVSVSDVPFPFSAQSGA	
N-19	VQLTLAFREGTINVDVETQFNQKTEAASRYMLTISDVSVSDVPFPFSAQSGA	
N-26	ASRYMLTISDVSVSDVPFPFSAQSGA	
N-30	ISDVSVSDVPFPFSAQSGA	
C-5	QFNQKTEAASRYMLTISDVSVSDVPFPFSAQSGA	
N-19/C-5	ASRYMLTISDVSVSDVPFPFSAQSGA	
N-10	QFNQKTEAASRYMLTISDVSVSDVPFPFSAQSGA	
C-10	GTINVDVETQFNQKTEAASRYMLTISDVSVSDVPFPFSAQSGA	
		TM

Fig. 78B

45C11-1	17H6-1	18B4-2	1E4-1	28F9-1	20A10-2
8A9-1	23H1-1	23H1-1	18G12-1	C3	C3
	32C1-1	32C1-1	25E6-1	E6	
	31A1-2	31A1-2	C3		
	39H3-2	39H3-2			
	3C5-2	3C5-2			

Fig. 78C

T47D	T47D	T47D
1500	1500	1500
H292		H292
H1975		H1975
SKOV-3	SKOV-3	SKOV-3
DUI145	DUI145	DUI145
HPAFII	HPAFII	HPAFII

Figure 78A-78C

FACS Analyses: Binding of antibodies to T47D breast cancer cells

Fig. 79A

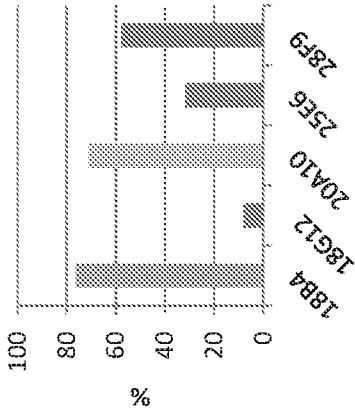


Fig. 79B

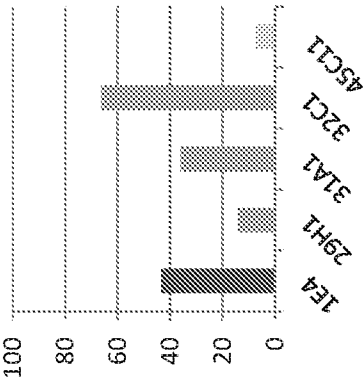


Fig. 79C

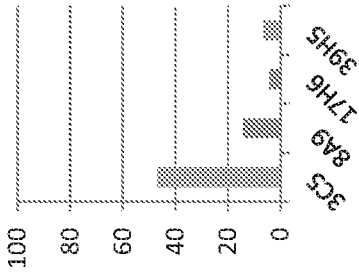


Fig. 79D

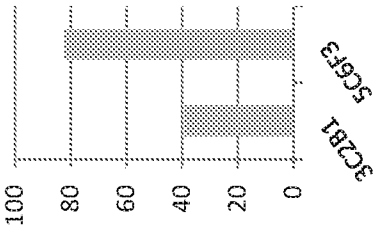


Fig. 79E

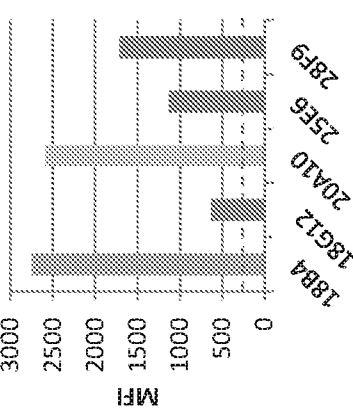


Fig. 79F

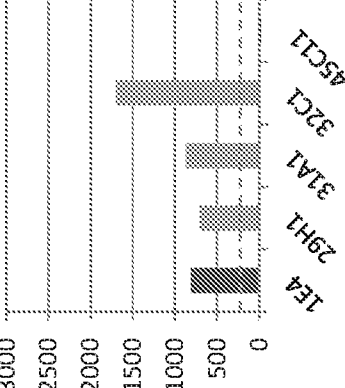


Fig. 79G

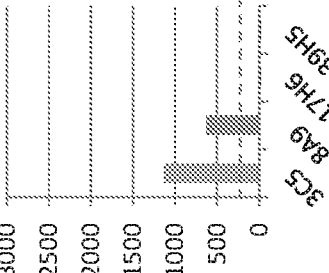


Fig. 79H

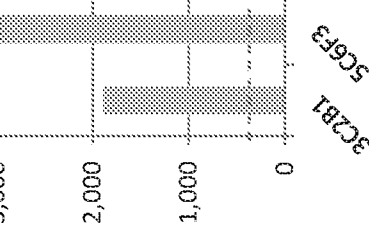


Fig. 79I

SNIKKRPQGVVQVLTARECTINVDVETIQNQKTEAAASKRYNLTISDVSVSDVPETESAQSGA

Figure 79A-79I

FACS Analyses: Binding of antibodies to 1500 breast cancer cells

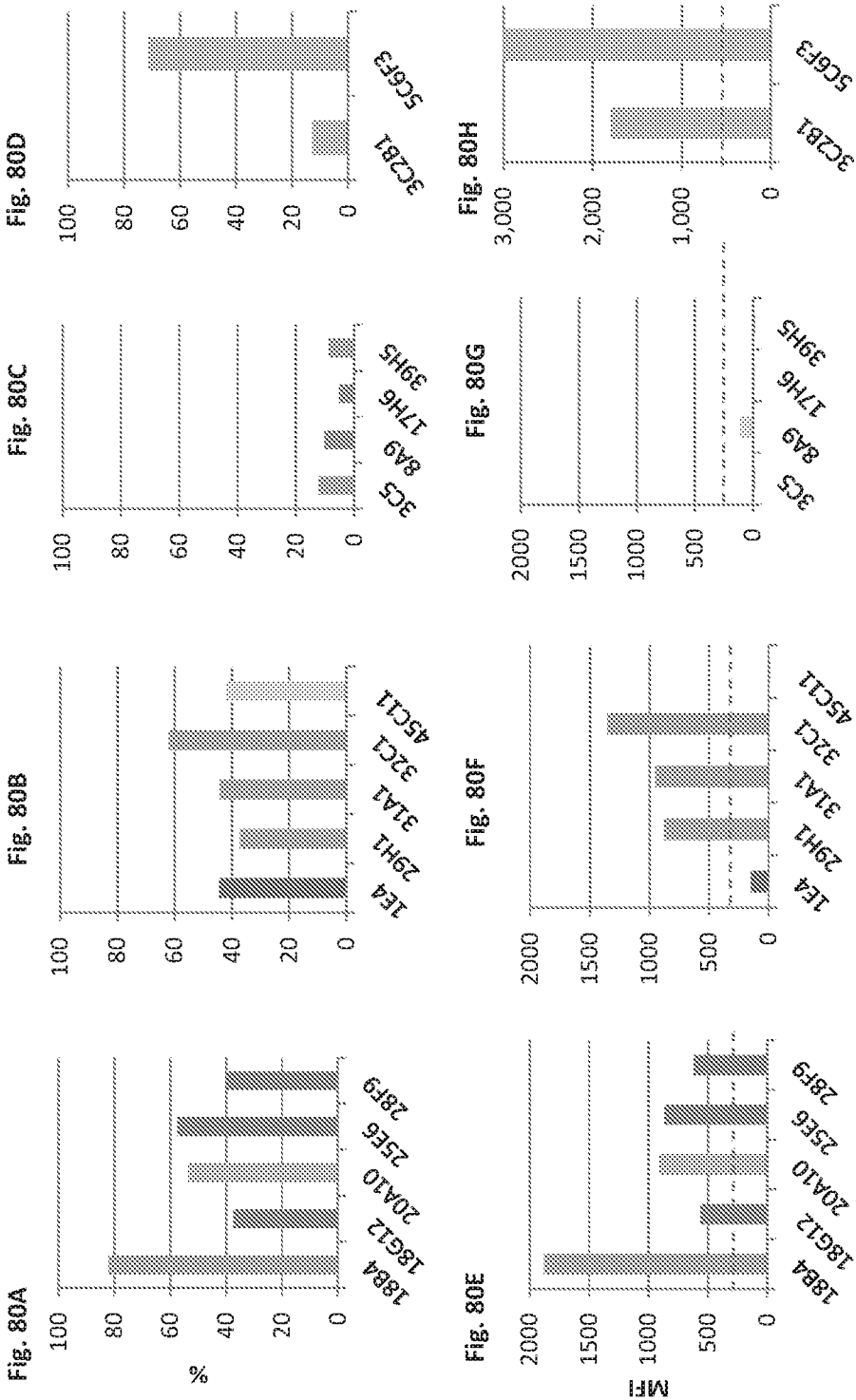


Fig. 80I  
SNINERPGSVVQVLTILAFREGTINVHDVETQFNQYKTEAASRYNLTISDVSVSVDVPTTFESAQSGA

Figure 80A-80I

FACS Analyses: Binding of antibodies to NCI-H292 lung cancer cells

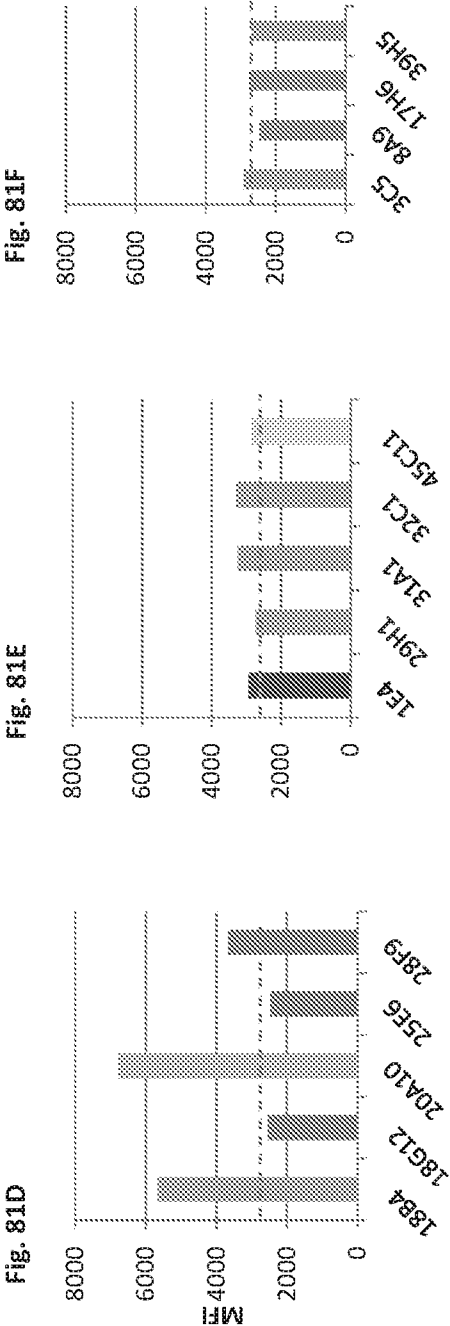
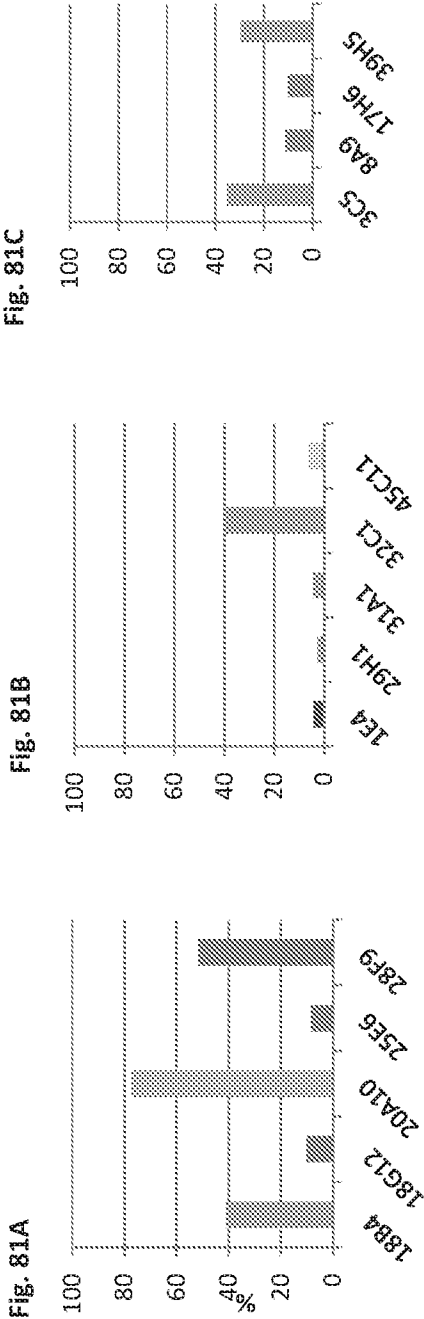
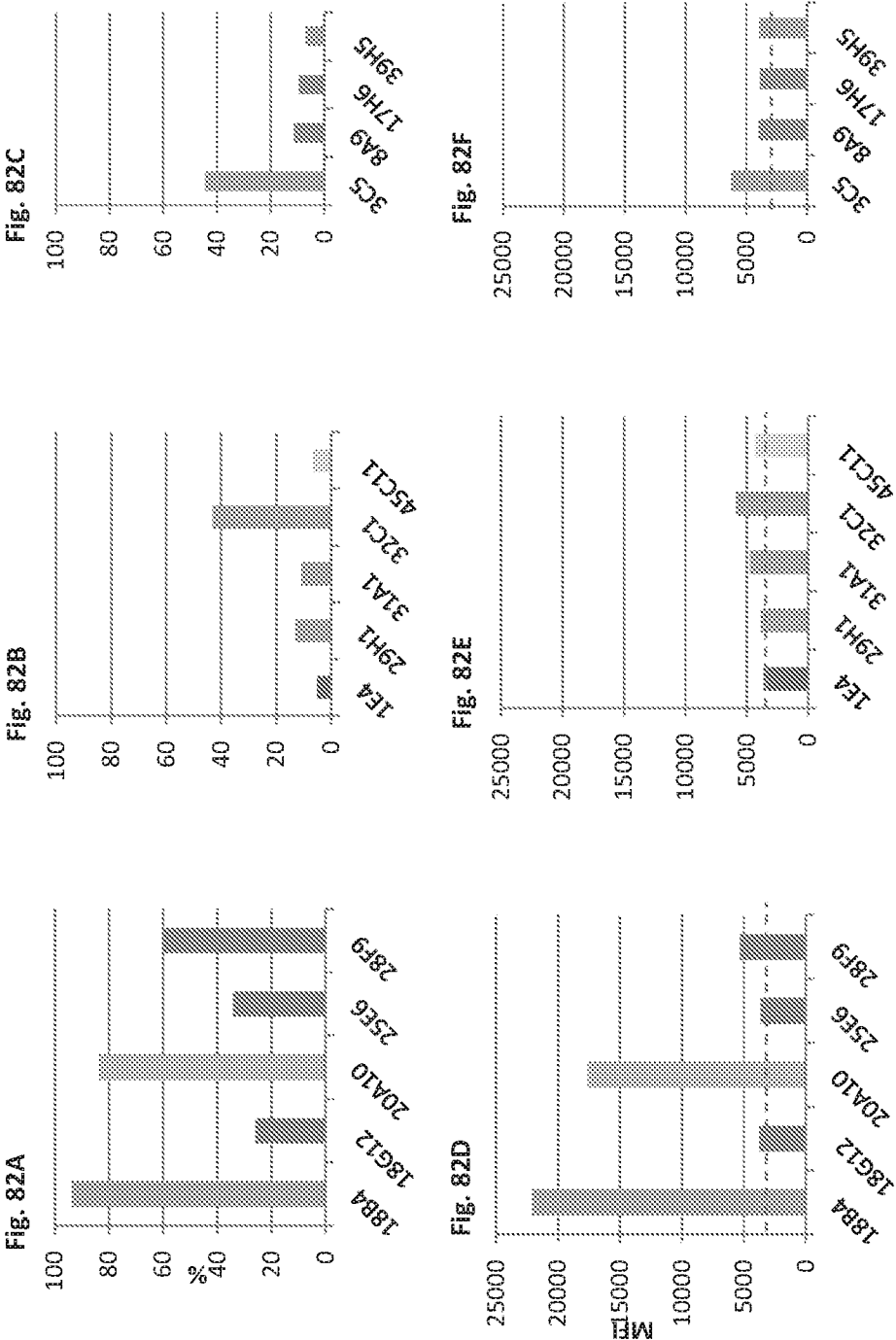


Fig. 81G

SNIKRPGSVVQLTLAIRECTINVHDVETQFNQYKTEAASRYNLTISDVSVDVPTPTSAQSGA

Figure 81A-81G

FACS Analyses: Binding of antibodies to NCI-H1975 lung cancer cells



**Fig. 82G**  
SNLILRRFGSVVQLILAFKREGTINVHDVETQENQYKTEAASRYNLLISDVSVDVPEPEEAQSGA  
Figure 82A-82G

FACS Analyses: Binding of antibodies to SKOV-3 ovarian cancer cells

Fig. 83A

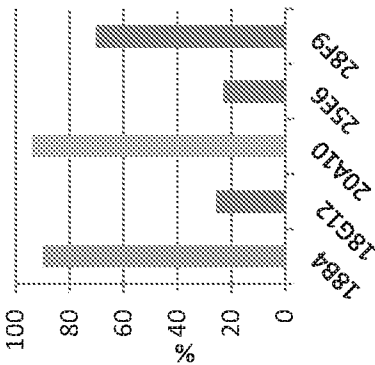


Fig. 83B

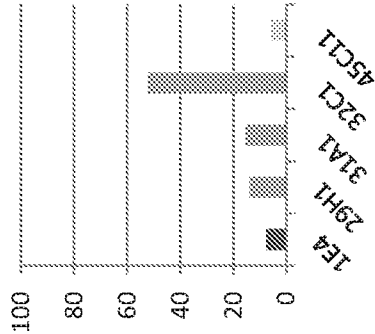


Fig. 83C

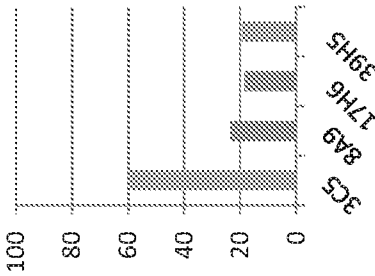


Fig. 83D

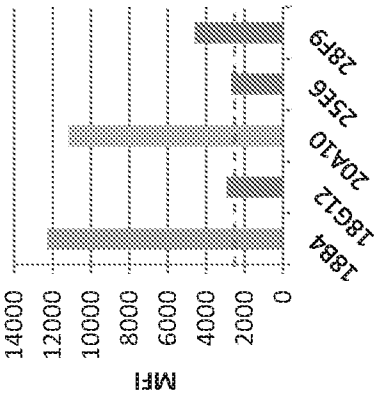


Fig. 83E

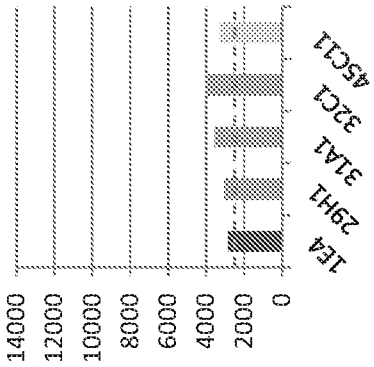


Fig. 83F

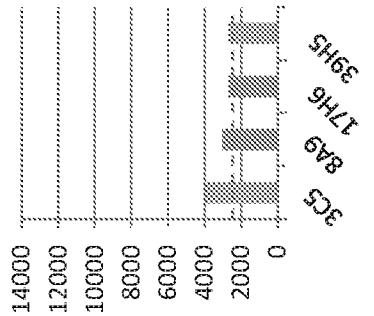


Fig. 83G

ENIKRRPGSVVQLTLAFREGTINVHDVETQENQYKTEAASKYNLTISDVSVSDVPFFPSAQSCA

Figure 83A-83G



FACS Analyses: Binding of antibodies to DU145 prostate cancer cells

Fig. 84A

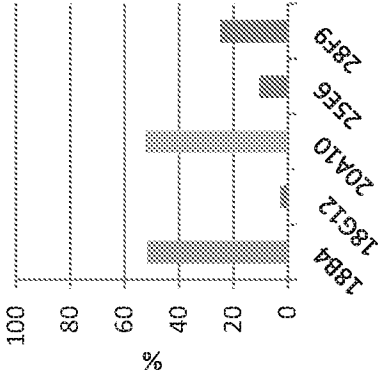


Fig. 84B

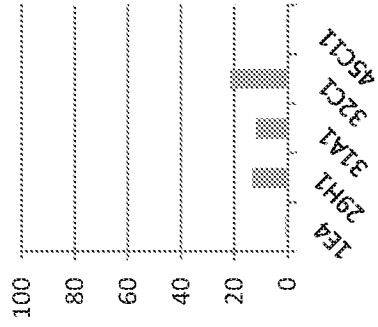


Fig. 84C

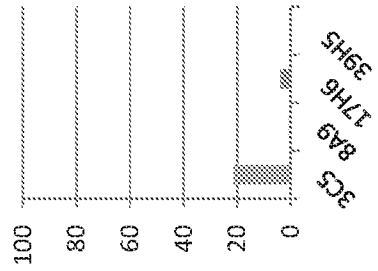


Fig. 84D

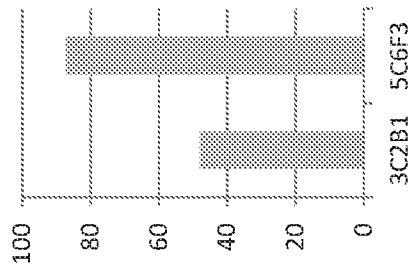


Fig. 84E

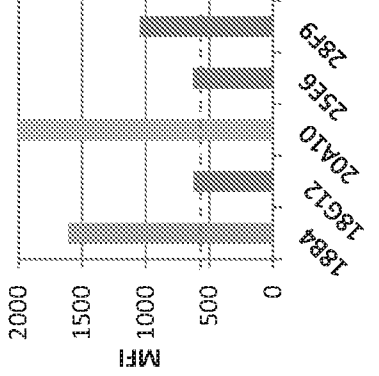


Fig. 84F

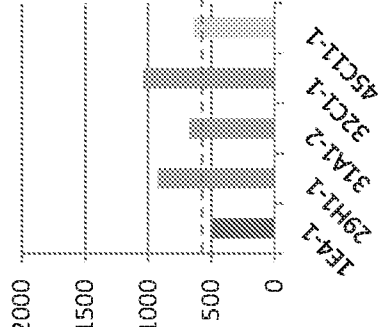


Fig. 84G

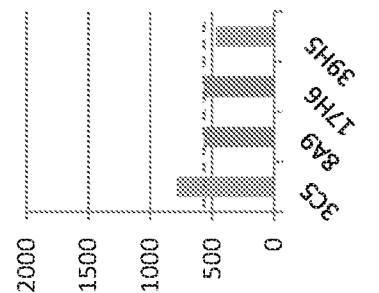


Fig. 84H

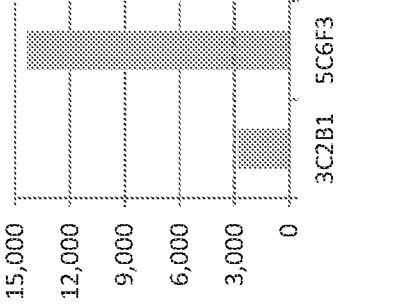


Fig. 84I

SNIKRPGGVVQLTAEKREGTINVDVETQFNQYKTEAASRYNLTISDVSVDVPEPEEAQSGA

Figure 84A-84I

FACS Analyses: Binding of antibodies to HPAF-II pancreatic cancer cells

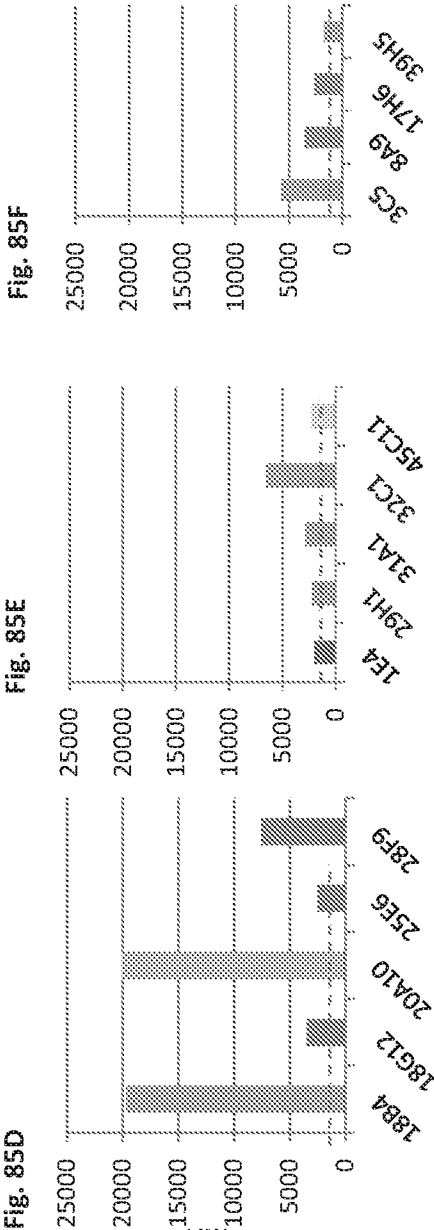
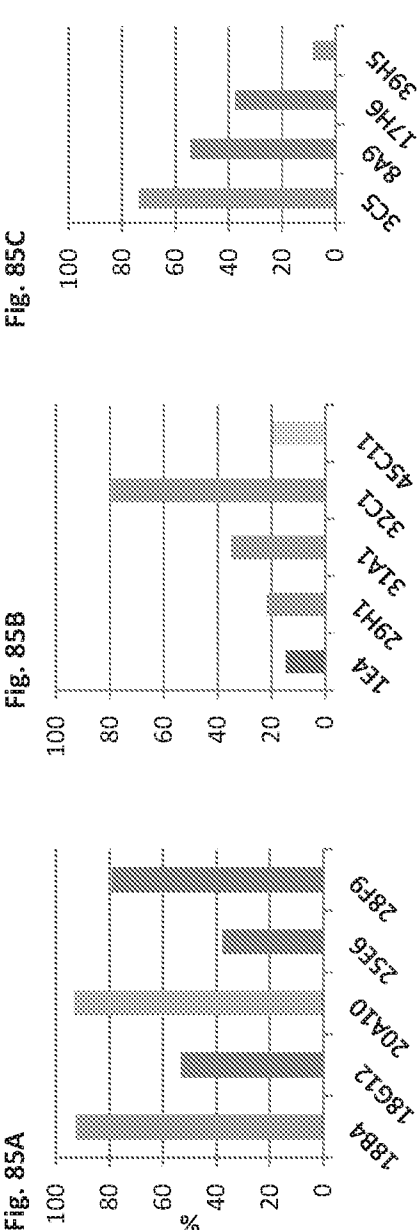


Fig. 85G

SNIKTRGGSVVQLTIAFREGTINVHDVETQFNQYKTEAASRYNLTISDVSVDVPEPFSAQSGA

Figure 85A-85G

FACS Analyses: Binding of antibodies to Capan-1 pancreatic cancer cells

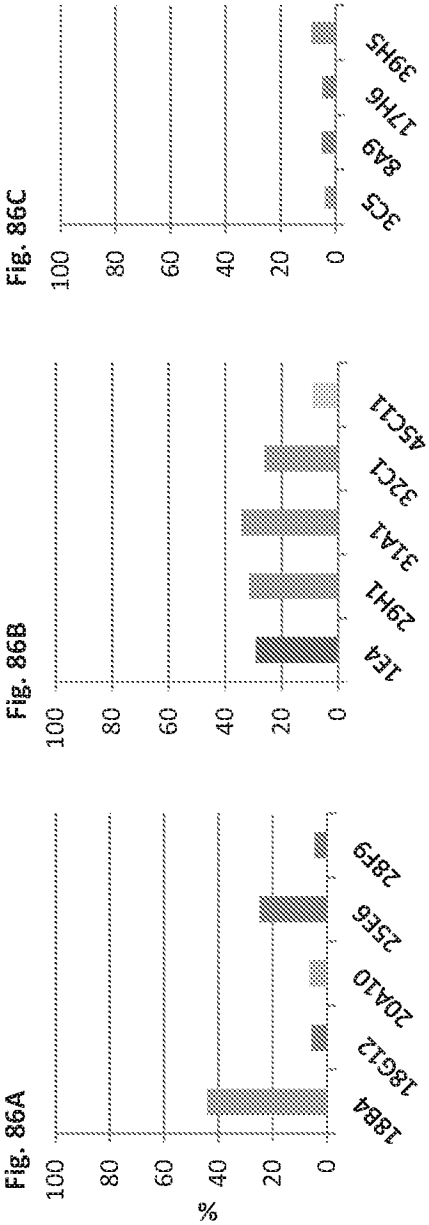


Fig. 86D

Fig. 86E

Fig. 86F

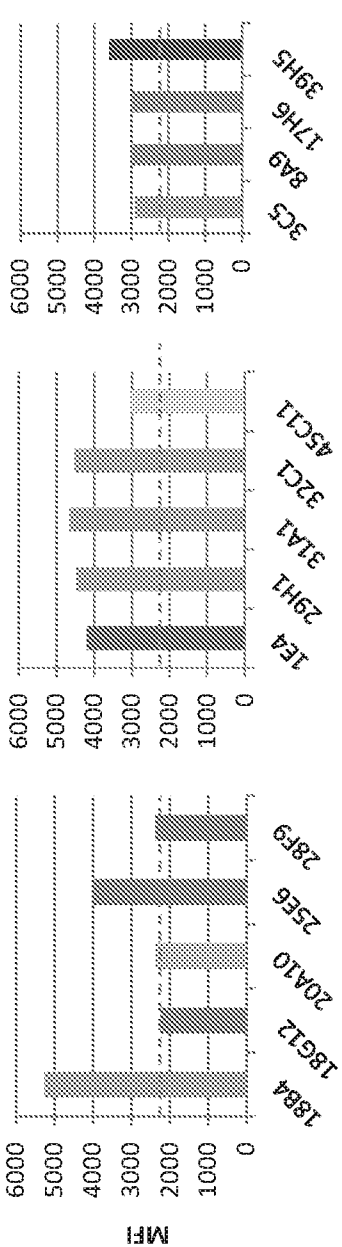


Fig. 86G

SNIKRPGSVVQLTLAFREGTINVDVETQENQYKTEAASRYNLTISDVSVDVPTPPSAQSGA

Figure 86A-86G

FACS Analyses: Binding of antibodies to MDA-MB-231 breast cancer cells

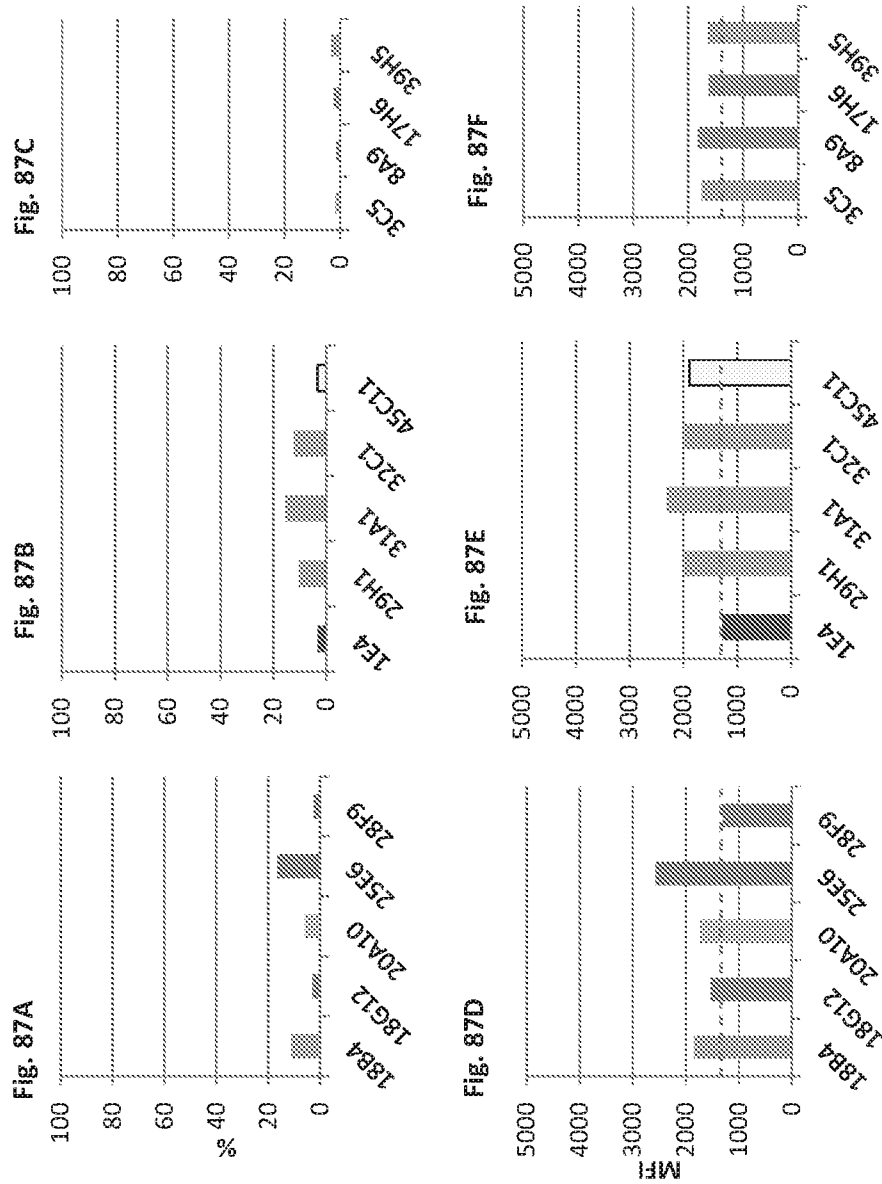
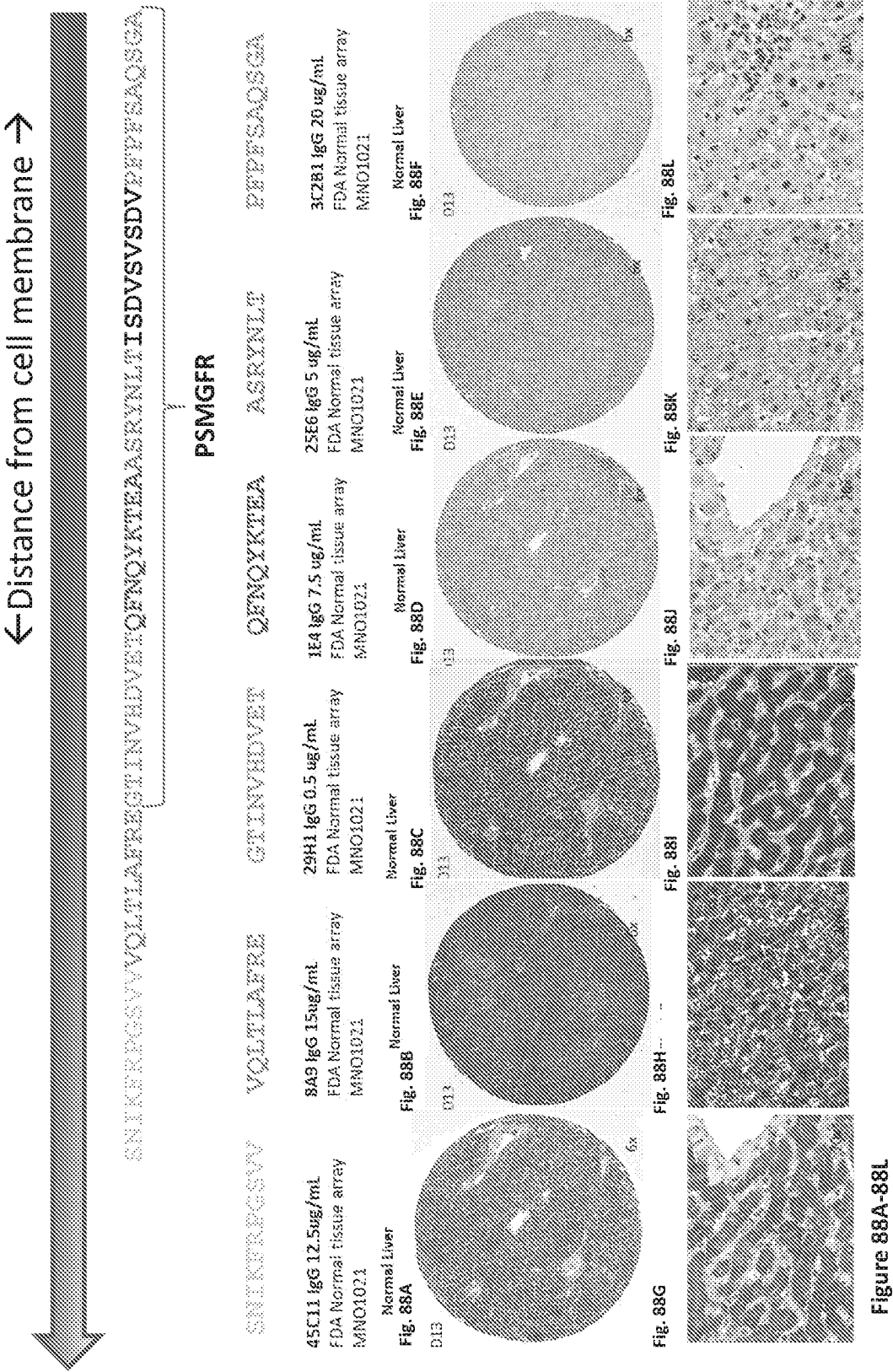


Fig. 87G

SNIKERPGSVVQVLTILARECTINVHDVETQENQYKTEAAQRNLIISDVSVDVPEPFSAQSGA

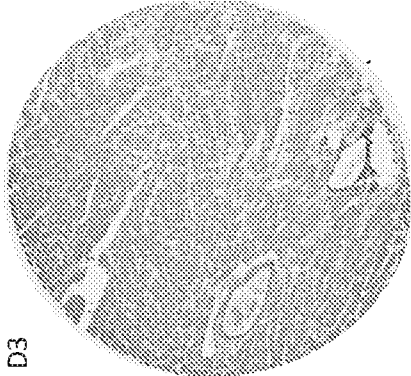
Figure 87A-87G



IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but not C-10: epitope comprises all or part of FPFS or FPFS is critical for antibody binding

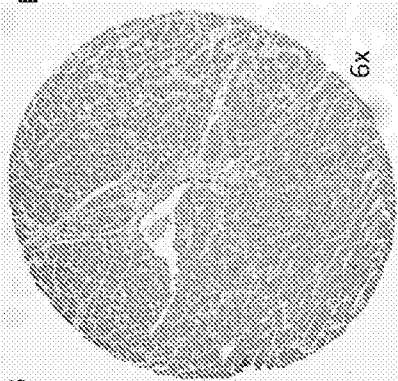
huMNC2-scFv 50 ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 89A



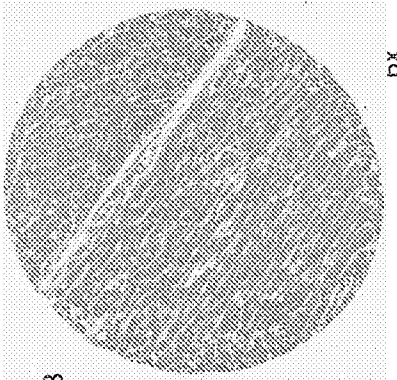
Mn-E6 IgG 2.50 ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 89B



20A10 IgG 0.25 ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 89C



3C2B1 IgG 20 ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 89D

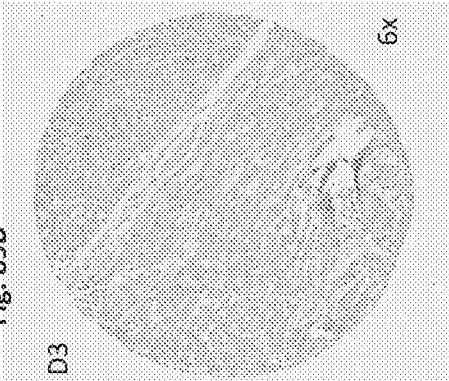


Fig. 89E

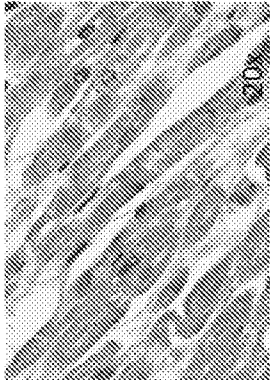


Fig. 89F

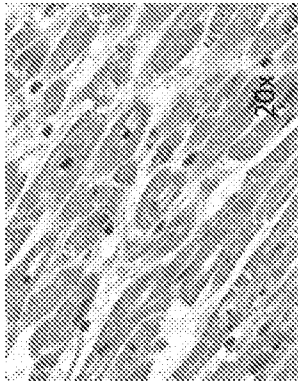


Fig. 89G

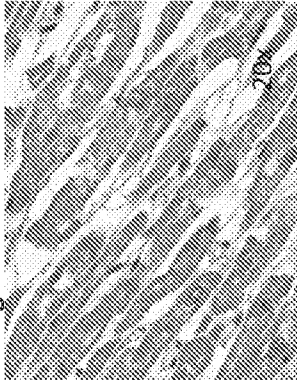


Fig. 89H

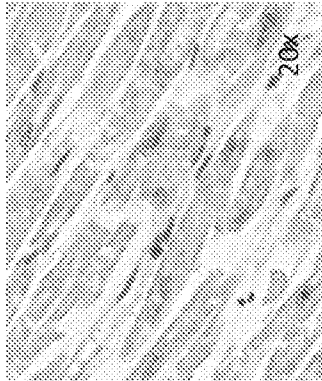
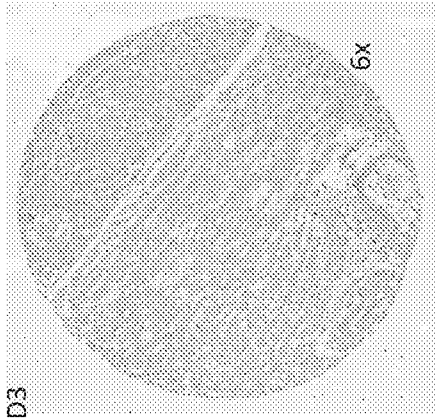


Figure 89A-89H

IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but also bind C-10: epitope comprises all or part of ASRYNLT

MNC3 IgG 5 ug/mL      25E6 IgG 5 ug/mL  
FDA Normal tissue array      FDA Normal tissue array  
MN88021      MN88021

Normal Heart  
Fig. 90A



Normal Heart  
Fig. 90B

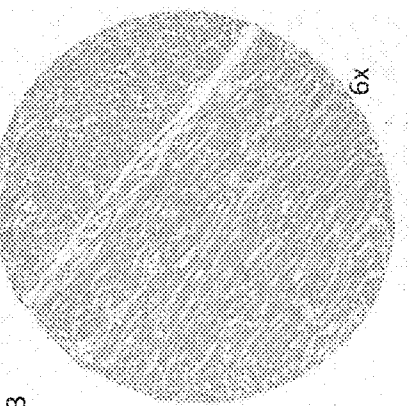


Fig. 90C

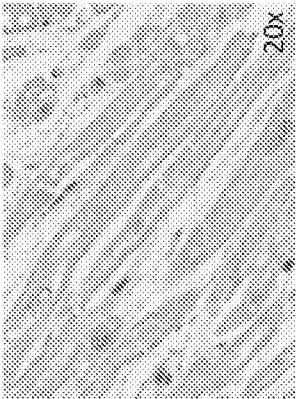


Fig. 90D

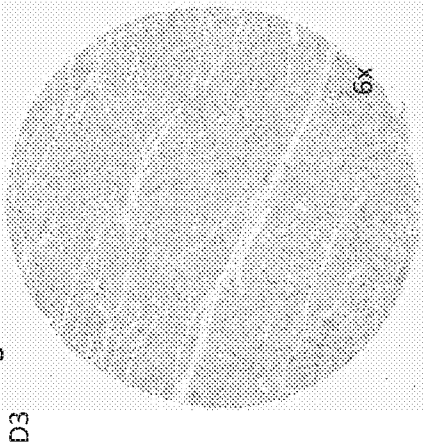


Figure 90A-90D

IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but also bind C-10: epitope comprises all or part of QFNQYKTEA

1E4 IgG 7.5 ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 91A



D3

Fig. 91B

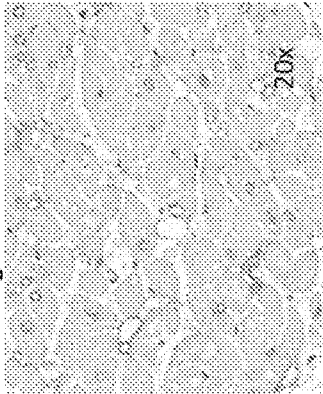


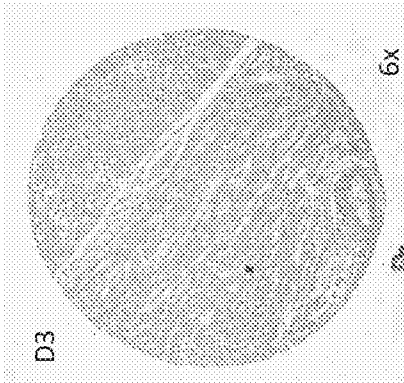
Figure 91A-91B



IHC of critical organs organized by antibody epitope: Antibodies that cannot bind to N-10 but can bind to C-10: epitope comprises all or part of GTINVHDVET

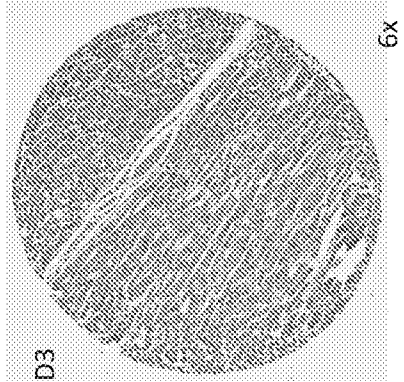
18B4 IgG 10ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 92A



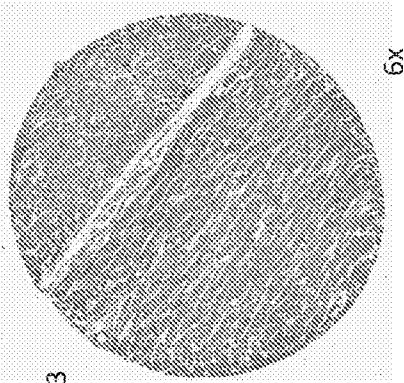
31A1 IgG 0.5 ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 92B



32C1 IgG 0.25 ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 92C



29H1 IgG 0.5 ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 92D

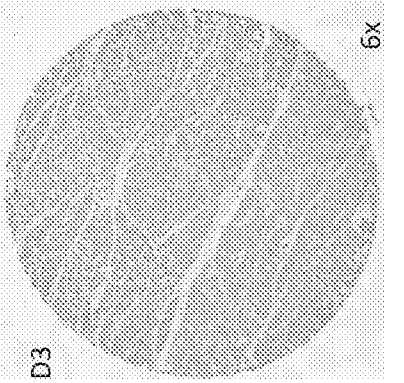


Fig. 92E

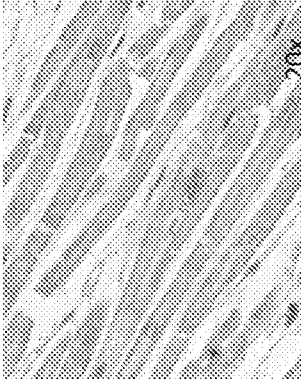


Fig. 92F

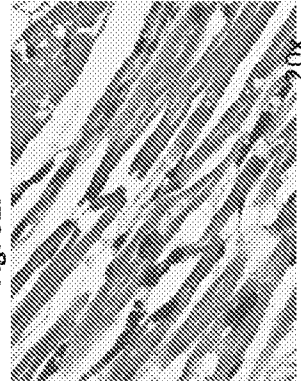


Fig. 92G

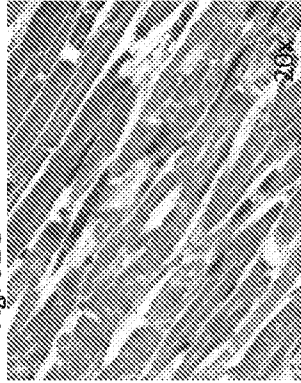


Fig. 92H

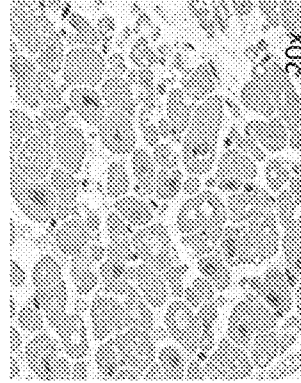


Figure 92A-92H

IHC of critical organs organized by antibody epitope: Antibodies that bind to sequences that are outside of, and N-terminal to, the PSMGR (MUC1\* extra cellular domain) sequence: epitope comprises all or part of VQLTLAFRE

8A9 IgG 15ug/mL	17H6 IgG 30 ug/mL
FDA Normal tissue array	FDA Normal tissue array
MN88021	MN88021

Normal Heart	Normal Heart
Fig. 93A	Fig. 93B

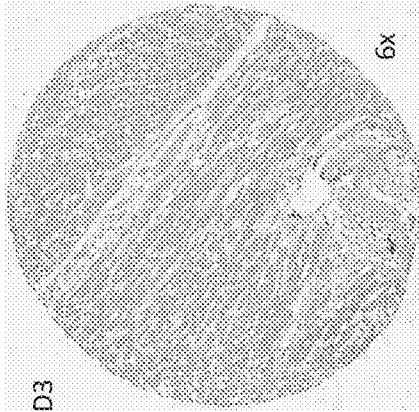
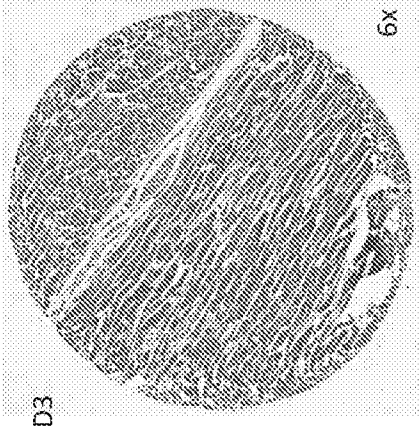


Fig. 93C

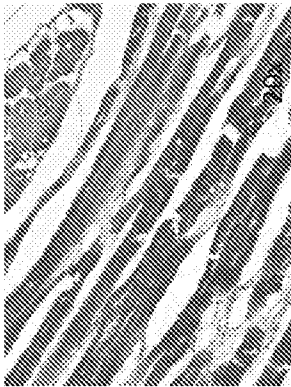


Fig. 93D

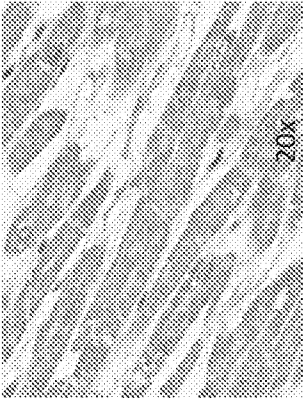
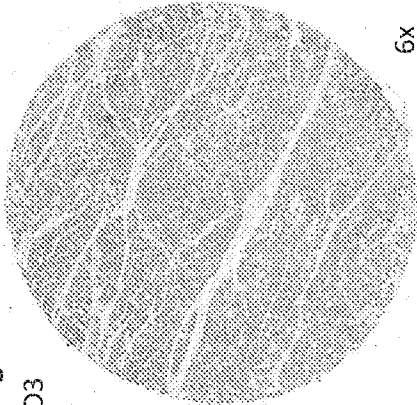


Figure 93A-93D

IHC of critical organs organized by antibody epitope: Antibodies that bind to sequences that are outside of, and N-terminal to, the PSMGR (MUC1\* extra cellular domain) sequence: epitope comprises all or part of SNIKFRPGSVV

45C11 IgG 12.5ug/mL  
FDA Normal tissue array  
MN88021

Normal Heart  
Fig. 94A



6x

Fig. 94B

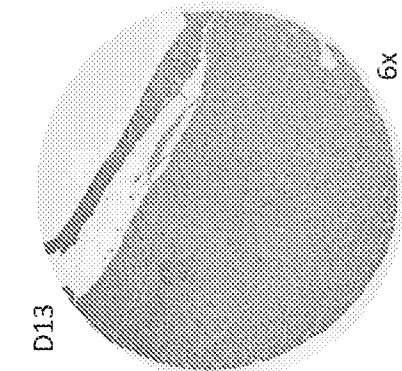


Figure 94A-94B

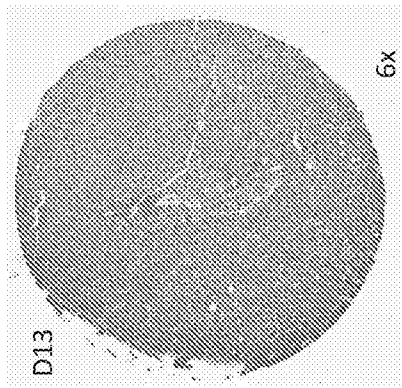
IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but not C-10: epitope comprises all or part of FPFS or FPFS is critical for antibody binding

huMNC2-scFv 50 ug/mL      MNE6 IgG 2.50 ug/mL      20A10 IgG 0.25 ug/mL      3C2B1 IgG 20 ug/mL  
FDA Normal tissue array      FDA Normal tissue array      FDA Normal tissue array      FDA Normal tissue array  
MN88021      MN88021      MN88021      MN88021

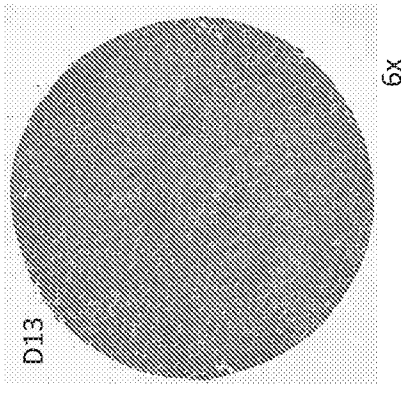
Normal Liver  
Fig. 95A



Normal Liver  
Fig. 95B



Normal Liver  
Fig. 95C



Normal Liver  
Fig. 95D

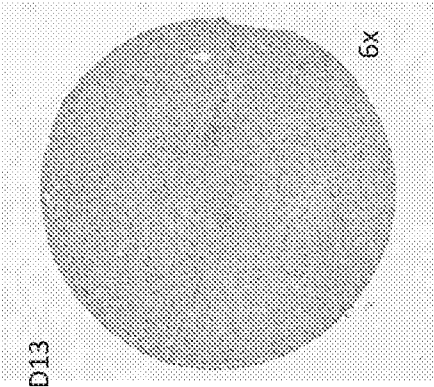


Fig. 95E

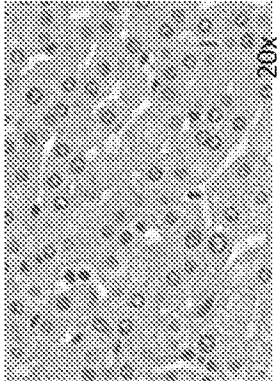


Fig. 95F

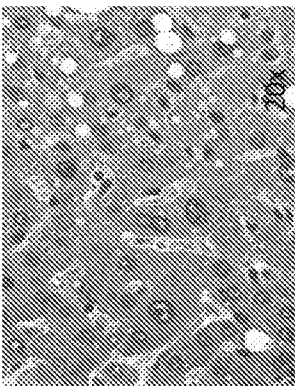


Fig. 95G

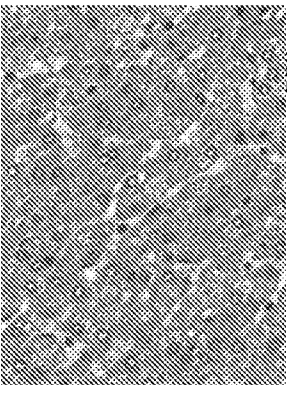


Fig. 95H

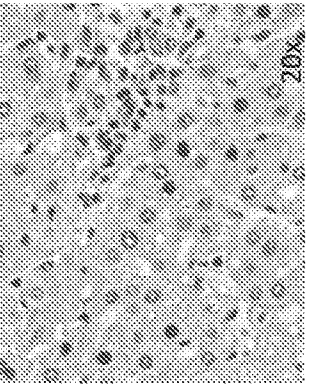


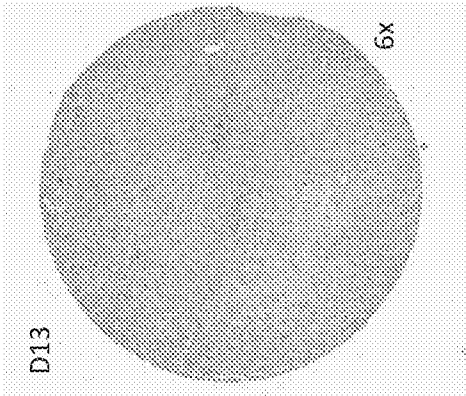
Figure 95A-95H

IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but also bind C-10: epitope comprises all or part of ASRYNLT

MNC3 IgG 5 ug/mL  
FDA Normal tissue array  
MN88021

25E6 IgG 5 ug/mL  
FDA Normal tissue array  
MN88021

Normal Liver  
Fig. 96A



Normal Liver  
Fig. 96B

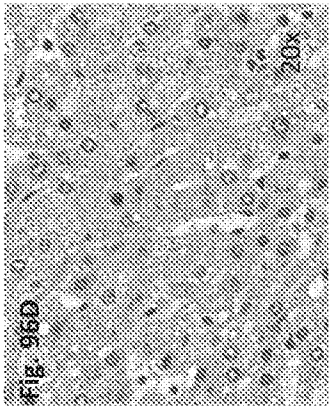
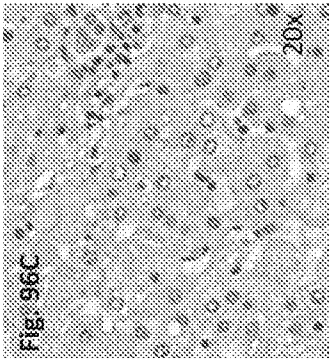
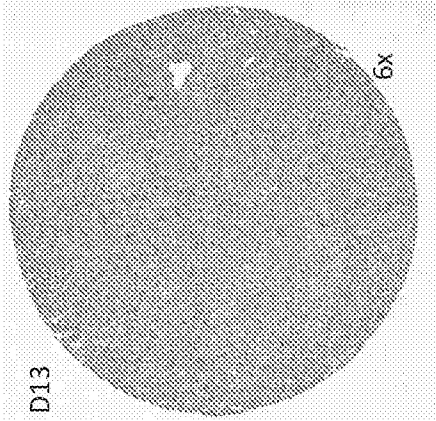


Figure 96A-96D

IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but also bind C-10: epitope comprises all or part of QFNQYKTEA

1E4 IgG 7.5 ug/mL  
FDA Normal tissue array  
MN88021  
Normal Liver  
Fig. 97A

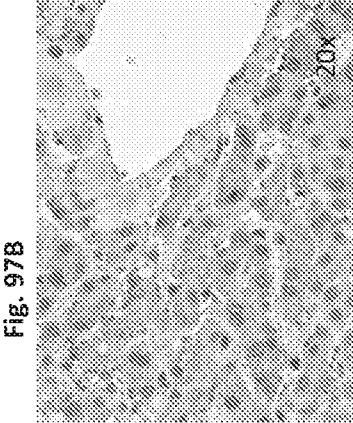
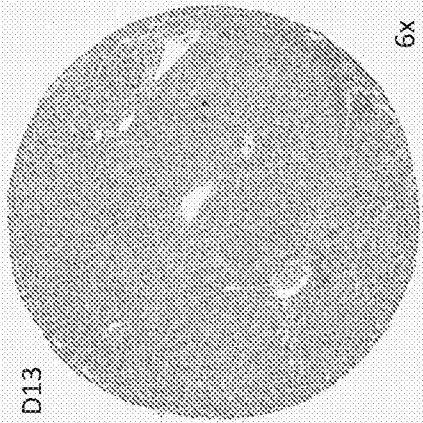


Figure 97A-97B

IHC of critical organs organized by antibody epitope: Antibodies that cannot bind to N-10 but can bind to C-10: epitope comprises all or part of GTINVHDVET

18B4 IgG 10ug/mL	31A1 IgG 0.5 ug/mL	32C1 IgG 0.25 ug/mL	29H1 IgG 0.5 ug/mL
FDA Normal tissue array	FDA Normal tissue array	FDA Normal tissue array	FDA Normal tissue array
MN88021	MN88021	MN88021	MN88021

Normal Liver  
Fig. 98A

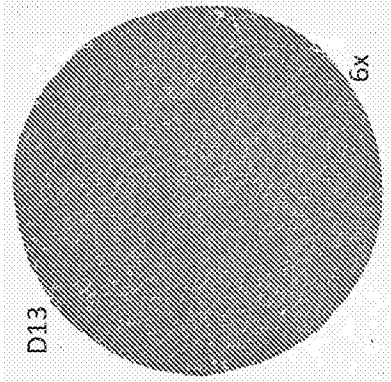
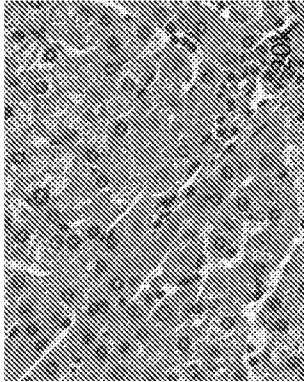


Fig. 98E



Normal Liver  
Fig. 98B

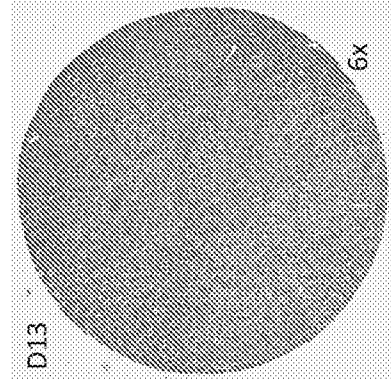
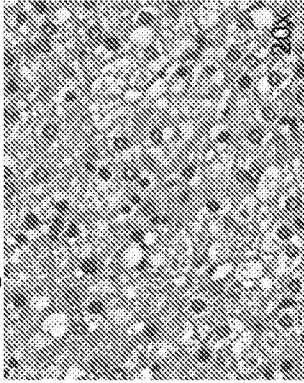


Fig. 98F



Normal Liver  
Fig. 98C

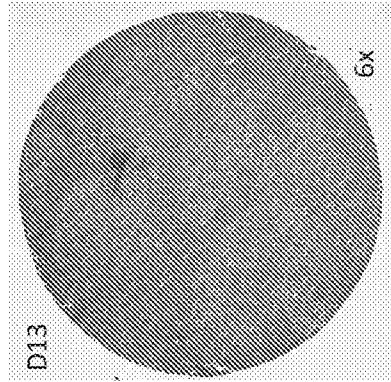
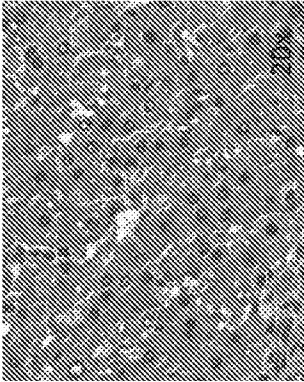


Fig. 98G



Normal Liver  
Fig. 98D

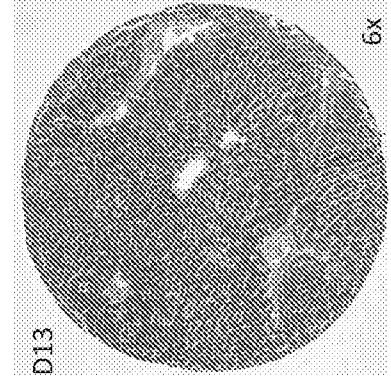


Fig. 98H

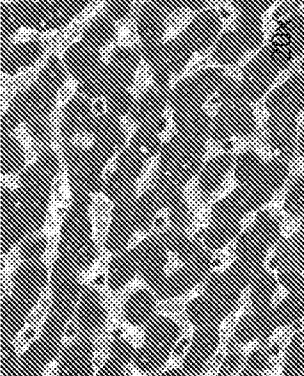


Figure 98A-98H



IHC of critical organs organized by antibody epitope: Antibodies that bind to sequences that are outside of, and N-terminal to, the PSMGFR (MUC1\* extra cellular domain) sequence: epitope comprises all or part of VQLTLAFRE

8A9 IgG 15ug/mL	17H6 IgG 30 ug/mL
FDA Normal tissue array	FDA Normal tissue array
MN88021	MN88021
Normal Liver	Normal Liver
Fig. 99A	Fig. 99B

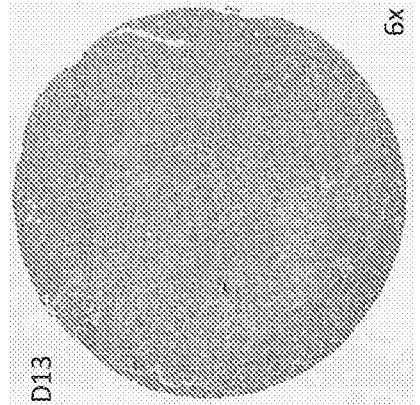
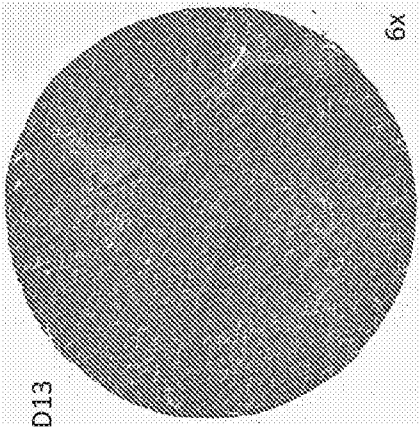


Fig. 99C

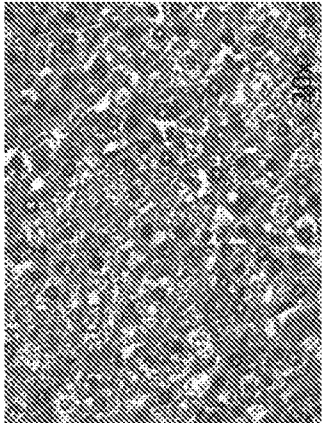


Fig. 99D

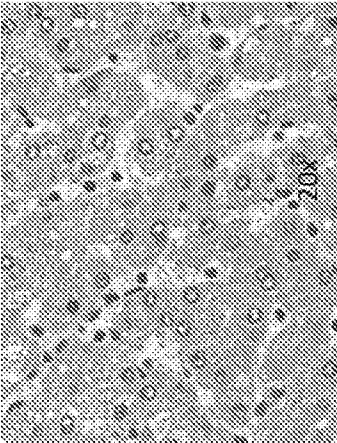


Figure 99A-99D



IHC of critical organs organized by antibody epitope: Antibodies that bind to sequences that are outside of, and N-terminal to, the PSMGR (MUC1\* extra cellular domain) sequence: epitope comprises all or part of SNIKFRPGSVV

45C11 IgG 12.5ug/mL  
FDA Normal tissue array  
MN88021

Normal Liver  
Fig. 100A

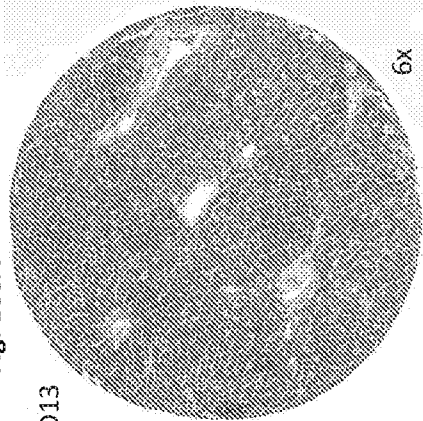


Fig. 100B

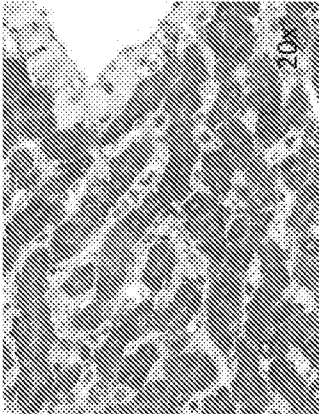


Figure 100A-100B

IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but not C-10: epitope comprises all or part of FPFS or FPFS is critical for antibody binding

huMNC2-scFv 50 ug/mL  
FDA Normal tissue array  
MN88021

Normal Lung  
Fig. 101A

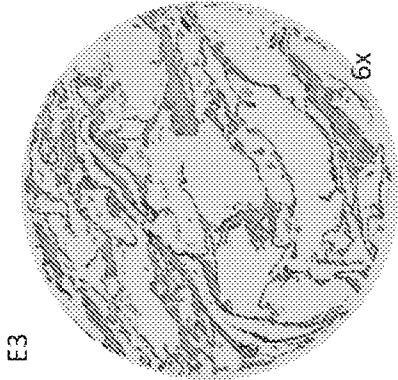


Fig. 101E



MNE6 IgG 2.50 ug/mL  
FDA Normal tissue array  
MN88021

Normal Lung  
Fig. 101B

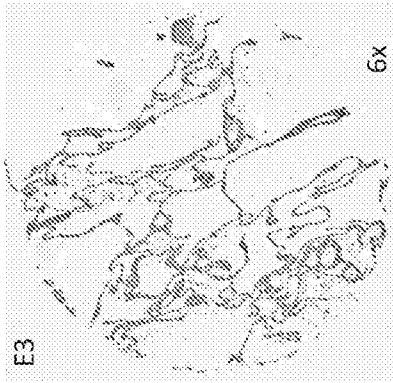


Fig. 101F



20A10 IgG 0.25 ug/mL  
FDA Normal tissue array  
MN88021

Normal Lung  
Fig. 101C

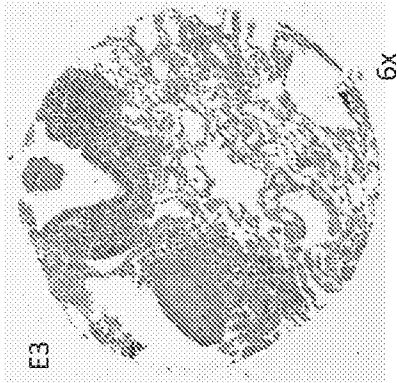
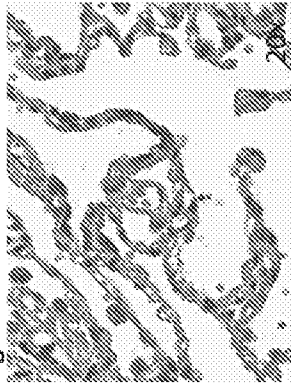


Fig. 101G



3C2B1 IgG 20 ug/mL  
FDA Normal tissue array  
MN88021

Normal Lung  
Fig. 101D

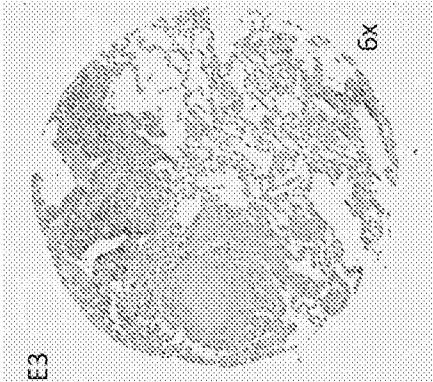


Fig. 101H

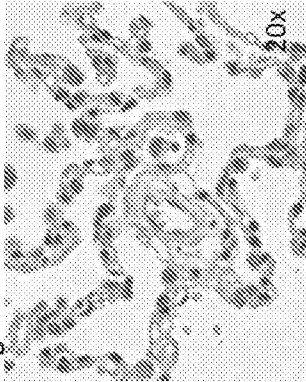


Figure 101A-101H

IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but also bind C-10: epitope comprises all or part of ASRYNLT

MNC3 IgG 5 ug/mL      25E6 IgG 5 ug/mL  
FDA Normal tissue array      FDA Normal tissue array  
MN88021      MN88021

Normal Lung      Normal Lung  
Fig. 102A      Fig. 102B

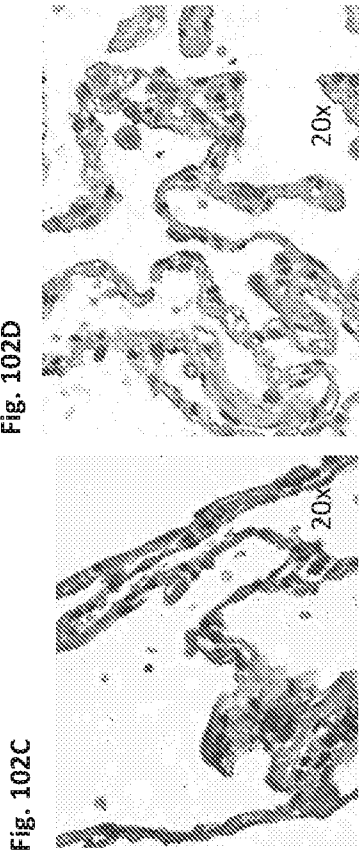
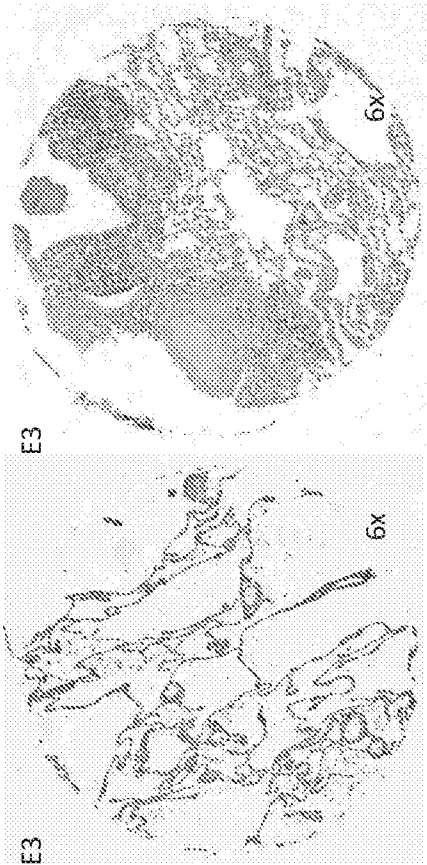


Figure 102A-102D

IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but also bind C-10: epitope comprises all or part of QFNQYKTEA

1E4 IgG 7.5 ug/mL  
FDA Normal tissue array  
MN88021  
Normal Lung  
Fig. 103A



Fig. 103B

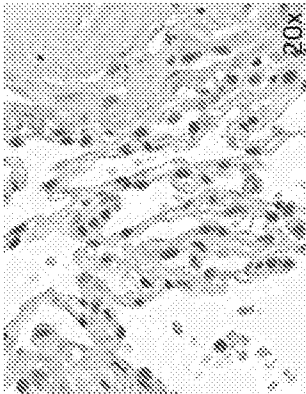


Figure 103A-103B

IHC of critical organs organized by antibody epitope: Antibodies that cannot bind to N-10 but can bind to C-10: epitope comprises all or part of GTINVHDVET

18B4 IgG 10ug/mL	31A1 IgG 0.5 ug/mL	32C1 IgG 0.25 ug/mL	29H1 IgG 0.5 ug/mL
FDA Normal tissue array	FDA Normal tissue array	FDA Normal tissue array	FDA Normal tissue array
MN88021	MN88021	MN88021	MN88021
Normal Lung	Normal Lung	Normal Lung	Normal Lung

Fig. 104A

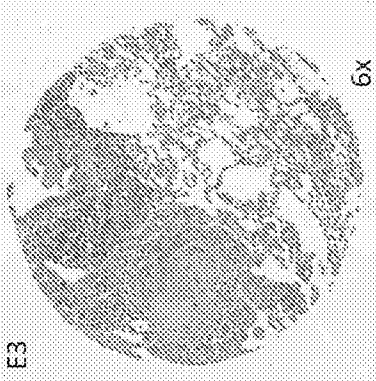


Fig. 104B

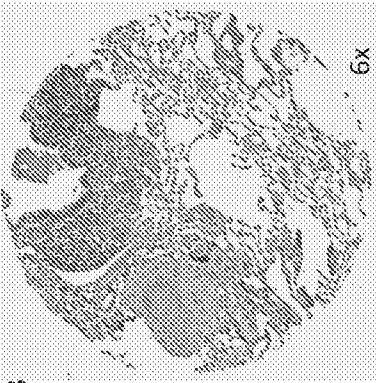


Fig. 104C

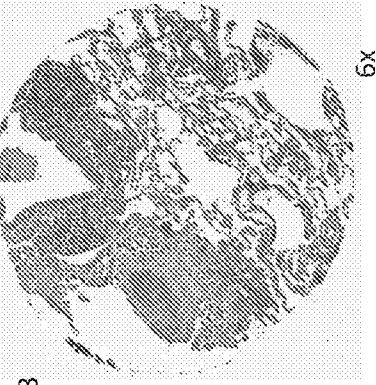


Fig. 104D

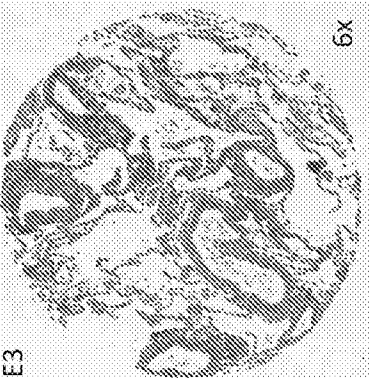


Fig. 104E

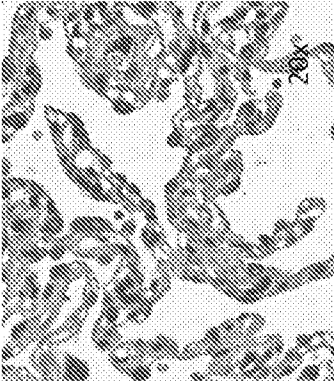


Fig. 104F

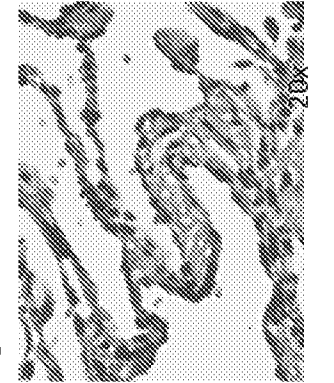


Fig. 104G

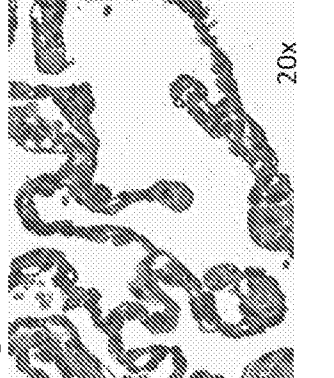


Fig. 104H

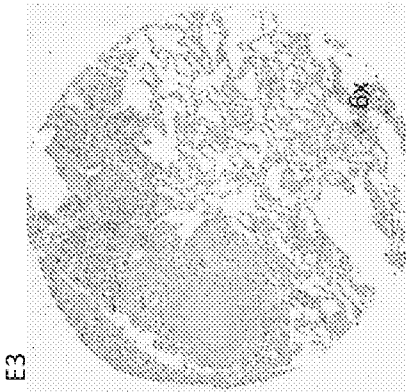
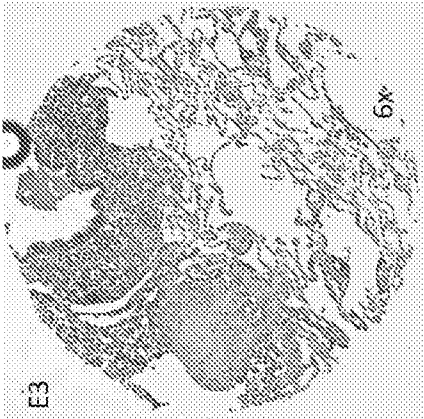


Figure 104A-104H

IHC of critical organs organized by antibody epitope: Antibodies that bind to sequences that are outside of, and N-terminal to, the PSMGFR (MUC1\* extra cellular domain) sequence: epitope comprises all or part of VQLTLAFRE

8A9 IgG 15ug/mL      17H6 IgG 30 ug/mL  
FDA Normal tissue array      FDA Normal tissue array  
MN88021      MN88021

Normal Lung  
Fig. 105A



Normal Lung  
Fig. 105B

Fig. 105C



Fig. 105D



Figure 105A-105D

IHC of critical organs organized by antibody epitope: Antibodies that bind to sequences that are outside of, and N-terminal to, the PSMGR (MUC1\* extra cellular domain) sequence: epitope comprises all or part of SNIKFRPGSVV

45C11 IgG 12.5ug/mL  
FDA Normal tissue array  
MN88021

Normal Lung  
Fig. 106A

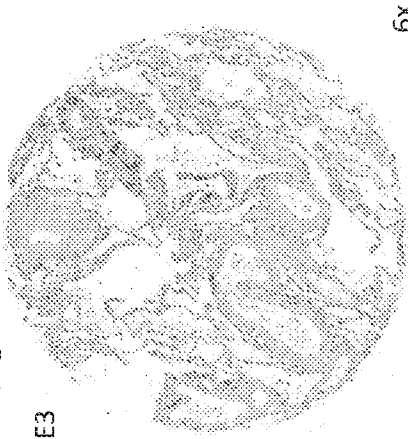


Fig. 106B

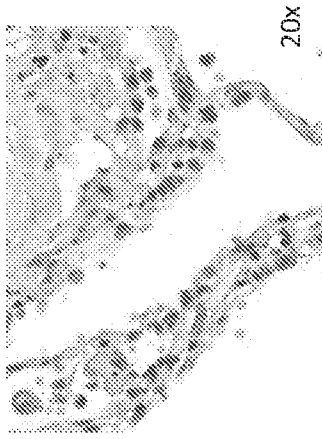


Figure 106A-106B

IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but not C-10: epitope comprises all or part of FPFS or FPFS is critical for antibody binding

huMNC2-scFv 50 ug/mL	MNE6 IgG 2.50 ug/mL	20A10 IgG 0.25 ug/mL	3C2B1 IgG 20 ug/mL
FDA Normal tissue array	FDA Normal tissue array	FDA Normal tissue array	FDA Normal tissue array
MN88021	MN88021	MN88021	MN88021

Normal Bone Marrow	Normal Bone Marrow	Normal Bone Marrow	Normal Bone Marrow
Fig. 107A	Fig. 107B	Fig. 107C	Fig. 107D

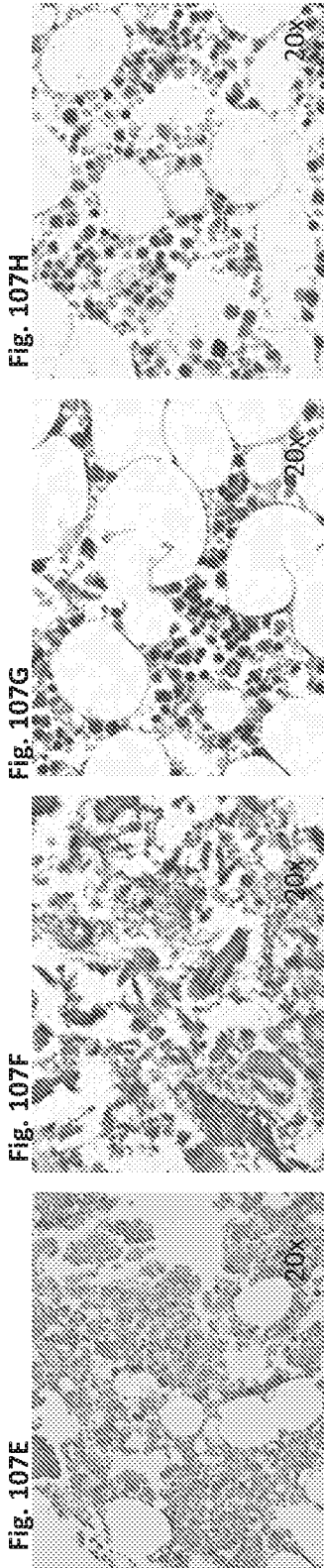
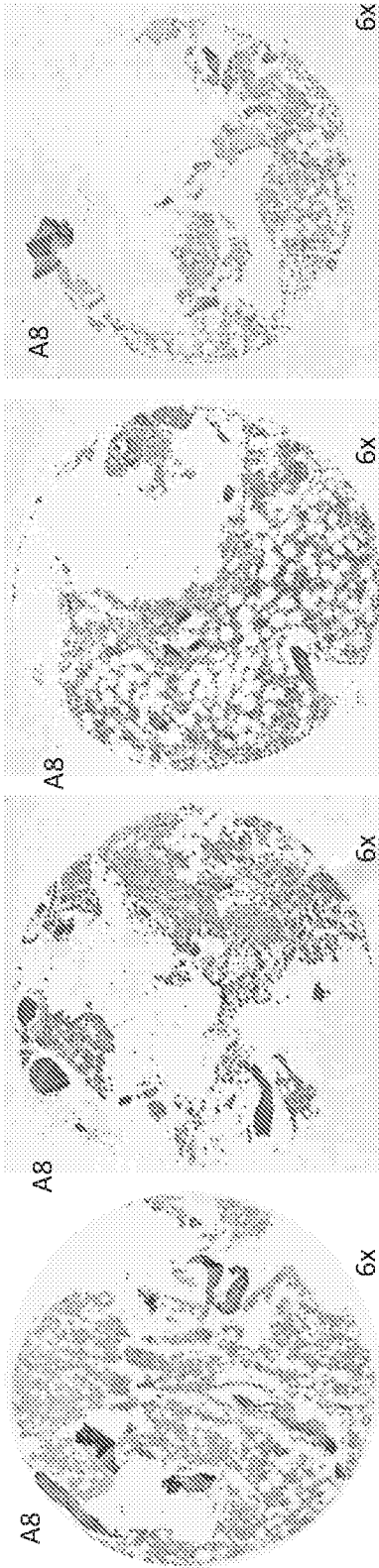


Figure 107A-107H



IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but also bind C-10: epitope comprises all or part of ASRYNLT

MNC3 IgG 5 ug/mL  
FDA Normal tissue array  
MN88021

25E6 IgG 5 ug/mL  
FDA Normal tissue array  
MN88021

Normal Bone Marrow  
Fig. 108A

Normal Bone Marrow  
Fig. 108B

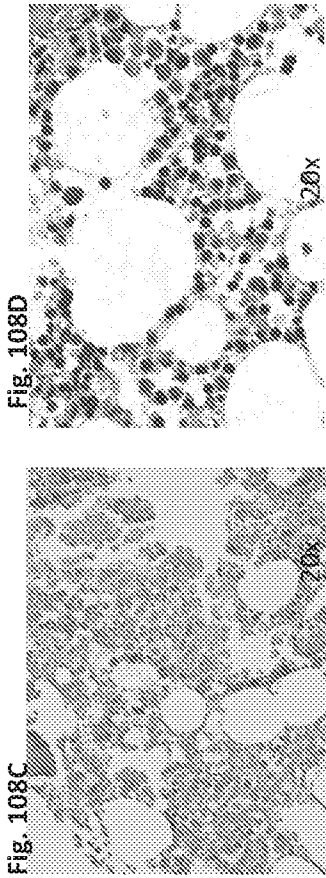
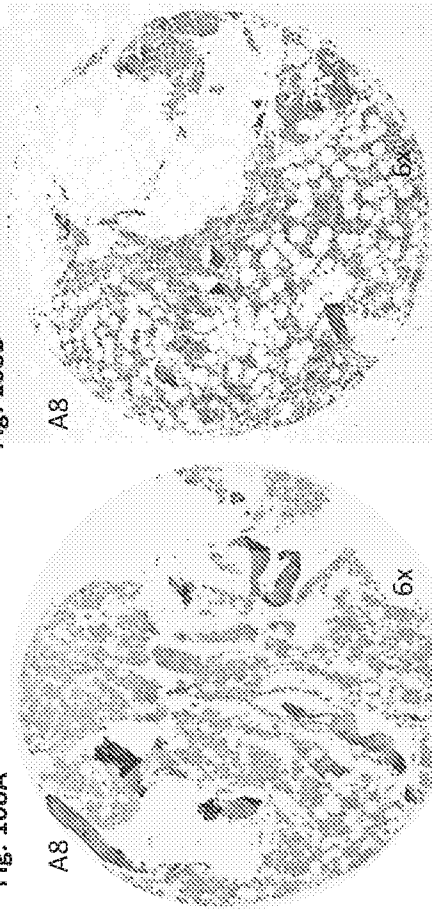


Figure 108A-108D

IHC of critical organs organized by antibody epitope: Antibodies that bind to N-10 but also bind C-10: epitope comprises all or part of QFNQYKTEA

1E4 IgG 7.5 ug/mL  
FDA Normal tissue array  
MN88021  
Normal Bone Marrow  
Fig. 109A

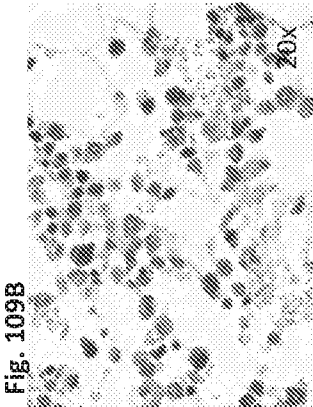
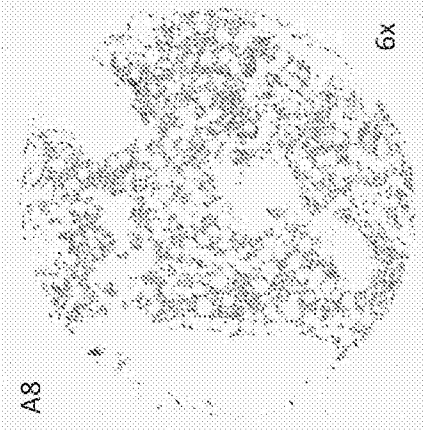


Figure 109A-109B

IHC of critical organs organized by antibody epitope: Antibodies that cannot bind to N-10 but can bind to C-10: epitope comprises all or part of GTINVHDVET

1884 IgG 10ug/mL	31A1 IgG 0.5 ug/mL	32C1 IgG 0.25 ug/mL	29H1 IgG 0.5 ug/mL
FDA Normal tissue array	FDA Normal tissue array	FDA Normal tissue array	FDA Normal tissue array
MN88021	MN88021	MN88021	MN88021
Normal Bone Marrow	Normal Bone Marrow	Normal Bone Marrow	Normal Bone Marrow

Fig. 110A



Fig. 110B

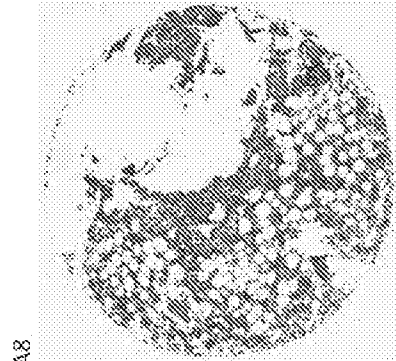


Fig. 110C

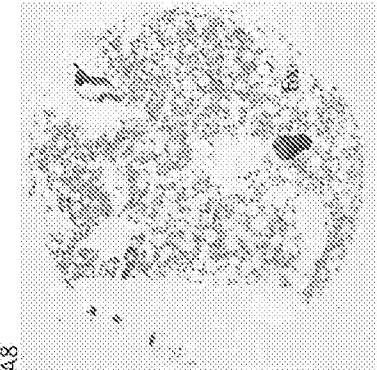


Fig. 110D

Fig. 110E



Fig. 110F

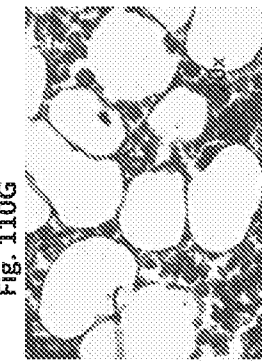


Fig. 110G

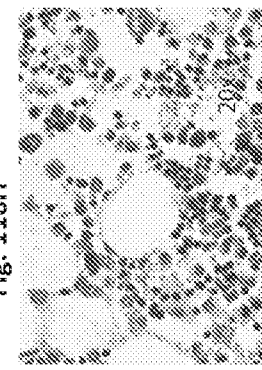


Fig. 110H

Figure 110A-110H

IHC of critical organs organized by antibody epitope: Antibodies that bind to sequences that are outside of, and N-terminal to, the PSMGFR (MUC1\* extra cellular domain) sequence: epitope comprises all or part of VQLTLAFRE

8A9 IgG 15ug/mL	17H6 IgG 30 ug/mL
FDA Normal tissue array	FDA Normal tissue array
MN88021	MN88021

Normal Bone Marrow      Normal Bone Marrow  
Fig. 111A      Fig. 111B

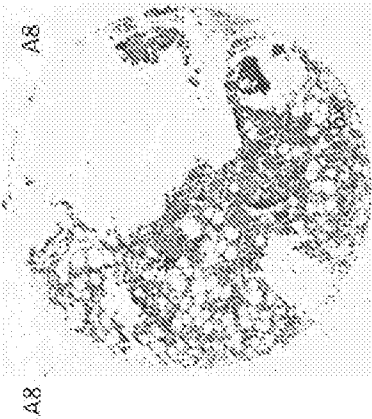


Fig. 111C

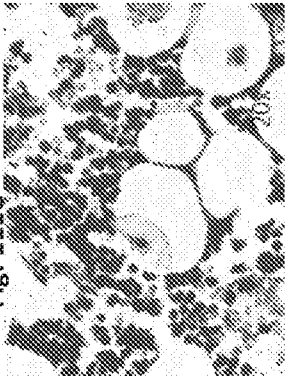


Fig. 111D

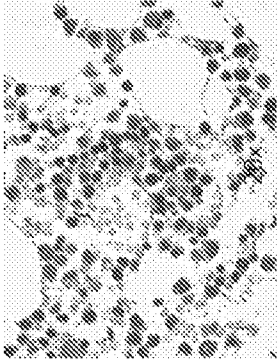


Figure 111A-111D

IHC of critical organs organized by antibody epitope: Antibodies that bind to sequences that are outside of, and N-terminal to, the PSMGR (MUC1\* extra cellular domain) sequence: epitope comprises all or part of SNIKFRPGSVV

45C11 IgG 12.5ug/mL  
FDA Normal tissue array  
MN88021

Normal Bone Marrow  
Fig. 112A

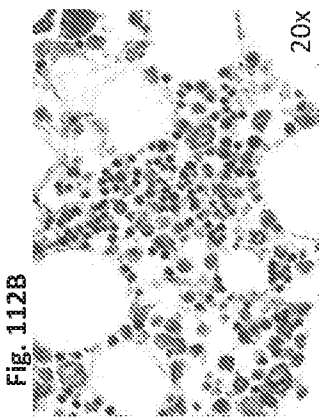
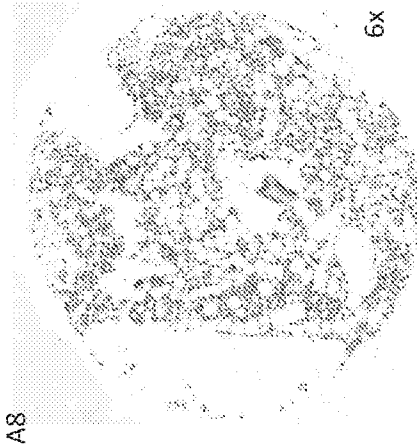


Figure 112A-112B

PSMGFR-antibody 20A10 IgG 0.25 ug/mL  
FDA Normal tissue array MNO1021

Fig. 113A

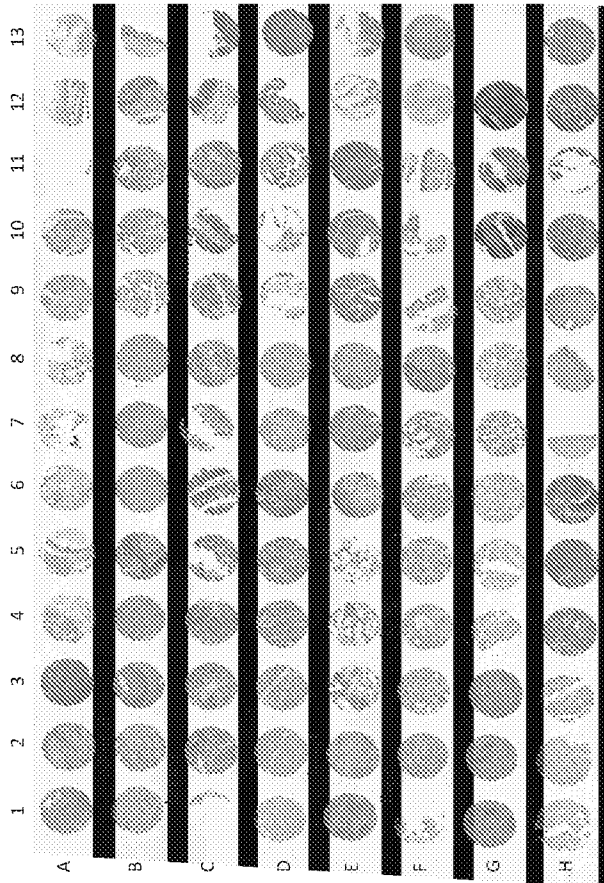


Fig. 113B

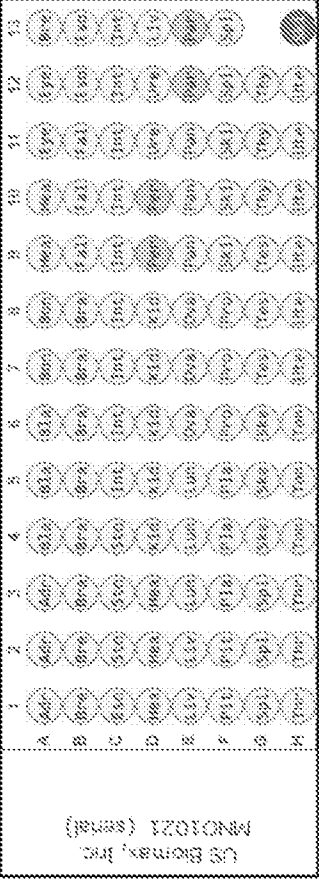


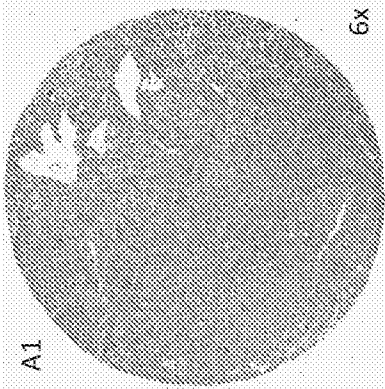
Figure 113A-113C

Fig. 113C

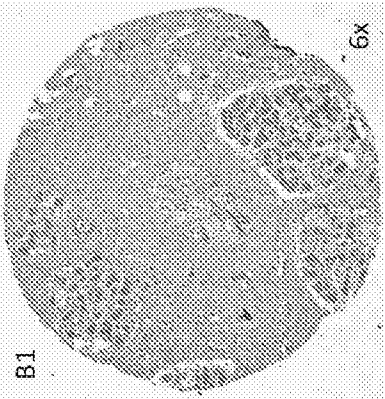
Position	Age	Sex	Organ/Anatomic Site	Pathology diagnosis	Position	Age	Sex	Organ/Anatomic Site	Pathology diagnosis
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	53	Adrenal gland	Normal	E3	M	72	Lung	Normal
A4	M	75	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	43	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bone, bone marrow	Normal	E7	F	46	Ovary	Normal
A8	M	43	Bone, bone marrow	Normal	E8	F	40	Ovary	Normal
A9	F	44	Head and neck, salivary gland	Normal	E9	M	42	Pancreas	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	26	Pancreas	Normal
A11	M	55	Eye	Normal	E11	F	59	Pancreas	Normal
A12	M	50	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	F	35	Breast	Normal	E13	M	72	Parathyroid	Adenoma
B1	F	38	Breast	Normal	F1	F	3	Pituitary gland	Normal
B2	F	33	Breast	Normal	F2	F		Pituitary gland	Normal
B3	M	58	Brain, cerebellum	Normal	F3	F	32	Placenta	Normal
B4	M	65	Brain, cerebellum	Normal	F4	F	30	Placenta	Normal
B5	F		Brain, cerebellum	Normal	F5	F	27	Placenta	Normal
B6	M	58	Brain, cerebral cortex	Normal	F6	M	64	Prostate	Normal
B7	M	65	Brain, cerebral cortex	Normal	F7	M	65	Prostate	Normal
B8	F		Brain, cerebral cortex	Normal	F8	M	65	Prostate	Normal
B9	F	42	Fallopian tube	Normal	F9	F	32	Skin	Normal
B10	F	32	Fallopian tube	Normal	F10	F	68	Skin	Normal
B11	F	24	Fallopian tube	Normal	F11	M	26	Skin	Normal
B12	M	45	Esophagus	Normal	F12	M	47	Spinal cord	Normal
B13	F	68	Esophagus	Normal	G1	M	56	Spinal cord	Normal
C1	M	54	Esophagus	Normal	G2	M	27	Spleen	Normal
C2	M	45	Stomach	Normal	G3	M	60	Spleen	Normal
C3	M	49	Stomach	Normal	G4	M	31	Spleen	Normal
C4	M	77	Stomach	Normal	G5	F	60	Skeletal muscle	Normal
C5	M	45	Intestine, small intestine	Normal	G6	F	49	Skeletal muscle	Normal
C6	F	75	Intestine, small intestine	Normal	G7	M	70	Skeletal muscle	Normal
C7	F	75	Intestine, small intestine	Normal	G8	M	43	Testis	Normal
C8	M	2	Intestine, colon	Normal	G9	M	30	Testis	Normal
C9	M	48	Intestine, colon	Normal	G10	M	77	Testis	Normal
C10	M	74	Intestine, colon	Normal	G11	M	15	Thymus	Normal
C11	F	47	Intestine, rectum	Normal	G12	M	28	Thymus	Normal
C12	F	75	Intestine, rectum	Normal	G13	F	9	Thymus	Normal
C13	M	86	Intestine, rectum	Normal	H1	F	28	Thyroid	Normal
D1	M	56	Heart	Normal	H2	F	37	Thyroid	Normal
D2	M	46	Heart	Normal	H3	F	51	Thyroid	Normal
D3	F	34	Heart	Normal	H4	M	48	Tonsil	Normal
D4	F	52	Kidney, cortex	Normal	H5	M	37	Tonsil	Normal
D5	M	23	Kidney, cortex	Normal	H6	M	13	Tonsil	Normal
D6	M	63	Kidney, cortex	Normal	H7	F	38	Uterus, cervix	Normal
D7	F	52	Kidney, medulla	Normal	H8	F	35	Uterus, cervix	Normal
D8	F	52	Kidney, medulla	Normal	H9	F	44	Uterus, cervix	Normal
D9	M	59	Peripheral nerve	Schwannoma	H10	F	38	Uterus, endometrium	Normal
D10	F	29	Peripheral nerve	Schwannoma	H11	F	41	Uterus, endometrium	Normal
D11	F	44	Ureter	Normal	H12	F	46	Uterus, endometrium	Normal
D12	F	42	Ureter	Normal	H13	M	58	Skin	Malignant melanoma (tissue marker)
D13	F	57	Liver	Normal					

**PSMGFR-antibody 20A10 IgG 0.25 ug/mL**  
FDA Normal tissue array MNO1021

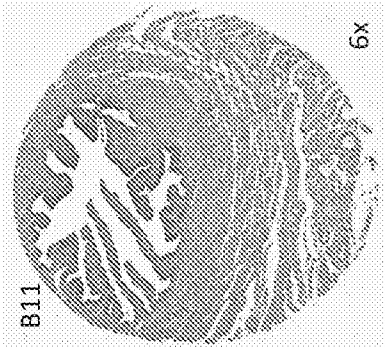
**Fig. 114A**  
Normal Adrenal Gland



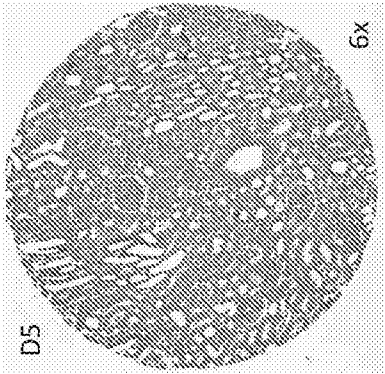
**Fig. 114B**  
Normal Breast



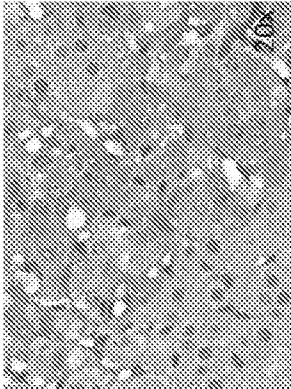
**Fig. 114C**  
Normal Fallopian Tubes



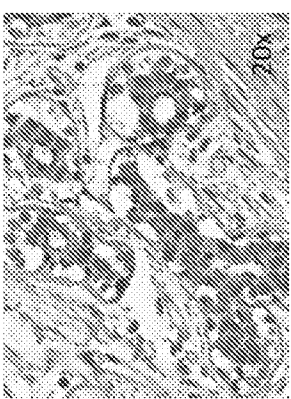
**Fig. 114D**  
Normal Kidney



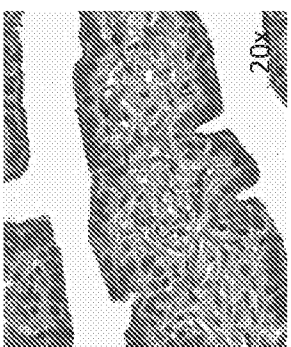
**Fig. 114E**



**Fig. 114F**



**Fig. 114G**



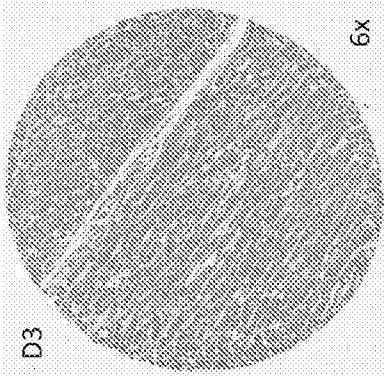
**Fig. 114H**



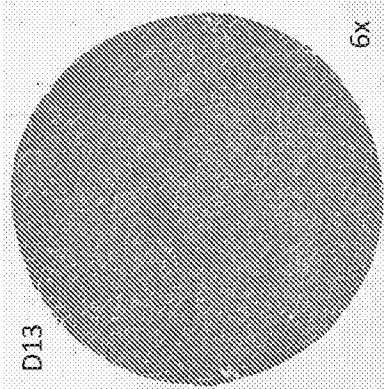
Figure 114A-114H

**PSMGFR-antibody 20A10 IgG 0.25 ug/mL**  
FDA Normal tissue array MNO1021

**Fig. 114I**  
Normal Heart



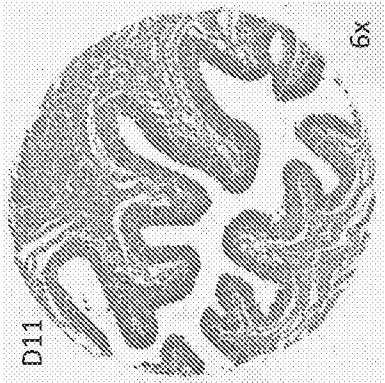
**Fig. 114J**  
Normal Liver



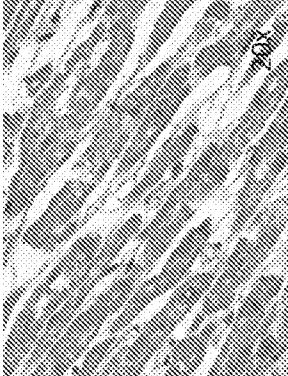
**Fig. 114K**  
Normal Lung



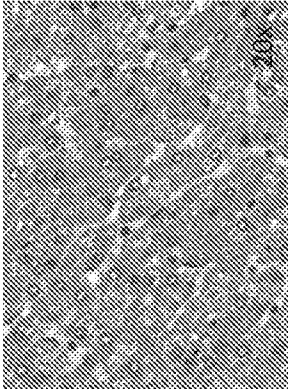
**Fig. 114L**  
Normal Ureter



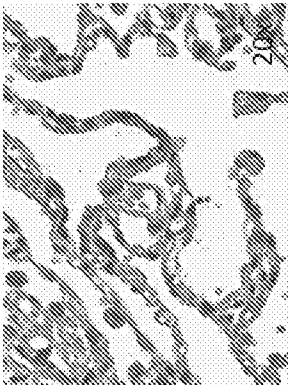
**Fig. 114M**



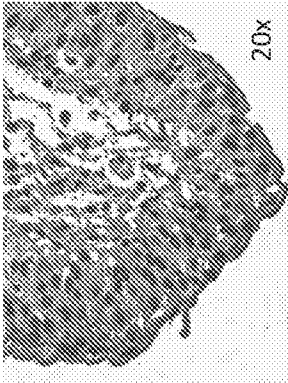
**Fig. 114N**



**Fig. 114O**



**Fig. 114P**

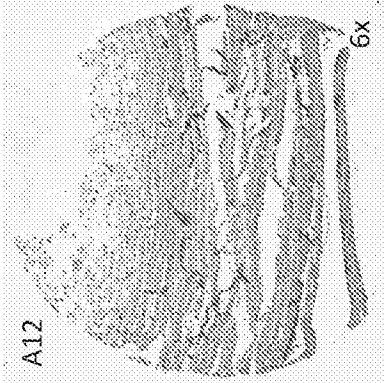


**Figure 114I-114P**

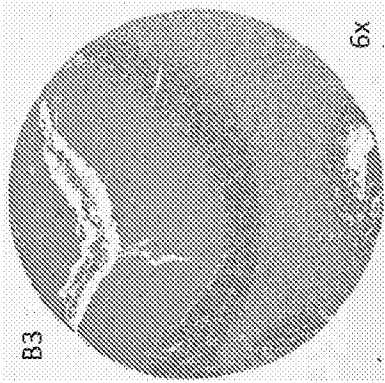


**PSMGFR-antibody 20A10 IgG 0.25 ug/mL**  
FDA Normal tissue array MNO1021

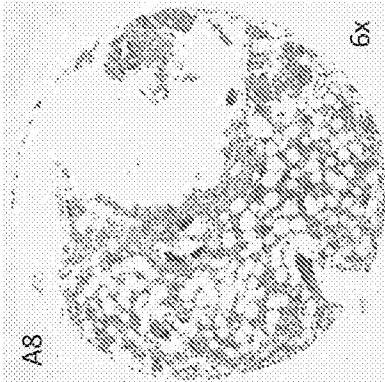
**Fig. 114Q**  
Normal Heart



**Fig. 114R**  
Normal Liver



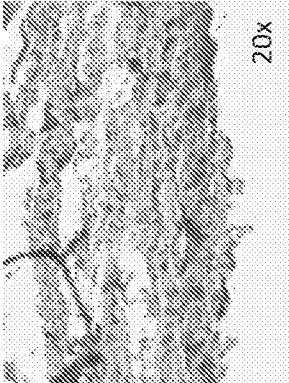
**Fig. 114S**  
Normal Lung



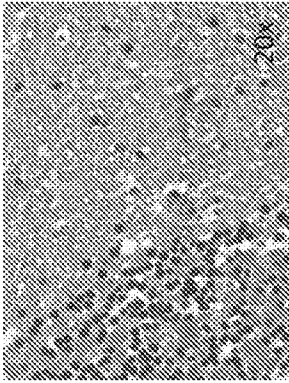
**Fig. 114T**  
Normal Ureter



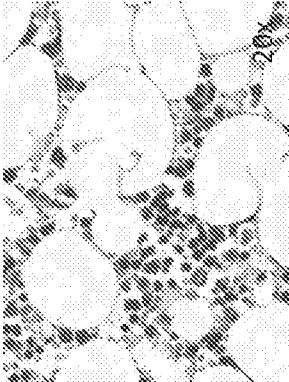
**Fig. 114U**



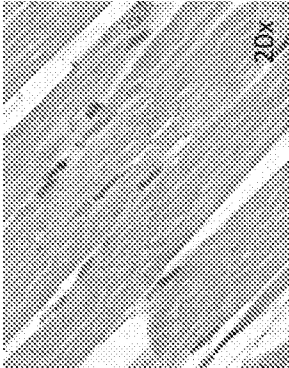
**Fig. 114V**



**Fig. 114W**



**Fig. 114X**



**Figure 114Q-114X**

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**PSM/GFR-antibody 20A10 IgG 0.25 ug/mL**  
Breast cancer tissue array BR1141

Fig. 115A

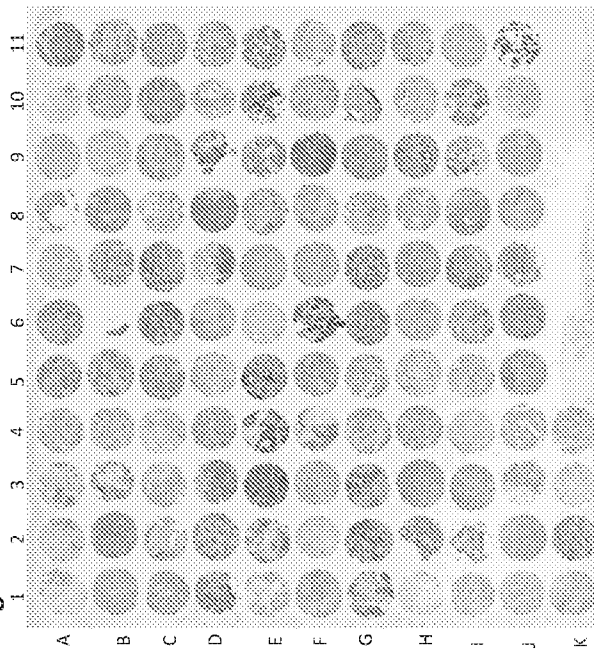


Fig. 115C

Position	Age	Sex	Primary diagnosis	TNM	Stage	Type	Position	Age	Sex	Primary diagnosis	TNM	Stage	Type
A1	45	F	Invasive ductal carcinoma	T2N0M0	2	IA	F1	52	F	Invasive ductal carcinoma	T2N0M0	2	IA
A2	38	F	Invasive ductal carcinoma	T2N0M0	2	IA	F2	44	F	Invasive ductal carcinoma	T2N0M0	2	IA
A3	42	F	Invasive ductal carcinoma	T2N0M0	2	IA	F3	35	F	Invasive ductal carcinoma	T2N0M0	2	IA
A4	40	F	Invasive ductal carcinoma	T2N0M0	2	IA	F4	38	F	Invasive ductal carcinoma	T2N0M0	2	IA
A5	70	F	Invasive ductal carcinoma	T2N1M0	2	IA	F5	83	F	Invasive ductal carcinoma	T2N1M0	2	IA
A6	48	F	Invasive ductal carcinoma	T2N0M0	2	IB	F6	59	F	Invasive ductal carcinoma	T2N1M0	2	IB
A7	42	F	Invasive ductal carcinoma	T2N1M0	2	IB	F7	43	F	Invasive ductal carcinoma	T2N0M0	2	IA
A8	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	F8	45	F	Invasive ductal carcinoma	T2N1M0	2	IA
A9	58	F	Invasive ductal carcinoma	T2N1M0	2	IB	F9	82	F	Invasive ductal carcinoma	T2N1M0	2	IB
A10	52	F	Invasive ductal carcinoma	T2N0M0	2	IB	F10	42	F	Invasive ductal carcinoma	T2N1M0	2	IB
A11	50	F	Invasive ductal carcinoma	T2N0M0	2	IA	F11	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
B1	43	F	Invasive ductal carcinoma	T2N0M0	2	IA	G1	45	F	Invasive ductal carcinoma	T2N0M0	2	IA
B2	37	F	Invasive ductal carcinoma	T2N1M0	2	IB	G3	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
B4	77	F	Invasive ductal carcinoma	T2N0M0	2	IA	G4	45	F	Invasive ductal carcinoma	T2N1M0	2	IB
B5	58	F	Invasive ductal carcinoma	T2N0M0	2	IA	G5	45	F	Invasive ductal carcinoma	T2N0M0	2	IA
B6	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	G6	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
B7	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	G7	53	F	Invasive ductal carcinoma	T2N0M0	2	IA
B8	40	F	Invasive ductal carcinoma	T2N0M0	2	IA	G8	42	F	Invasive ductal carcinoma	T2N1M0	2	IB
B9	53	F	Invasive ductal carcinoma	T2N0M0	2	IA	G9	38	F	Invasive ductal carcinoma	T2N1M0	2	IB
B10	47	F	Invasive ductal carcinoma	T2N1M0	2	IB	G10	46	F	Invasive ductal carcinoma	T2N1M0	2	IB
B11	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	H1	45	F	Invasive ductal carcinoma	T2N1M0	2	IB
C1	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	H2	53	F	Invasive ductal carcinoma	T2N0M0	2	IA
C2	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	H3	52	F	Invasive ductal carcinoma	T2N0M0	2	IB
C3	42	F	Invasive ductal carcinoma	T2N0M0	2	IA	H4	50	F	Invasive ductal carcinoma	T2N1M0	2	IB
C4	41	F	Invasive ductal carcinoma	T2N0M0	2	IA	H5	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
C5	58	F	Invasive ductal carcinoma	T2N0M0	2	IA	H6	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
C6	41	F	Invasive ductal carcinoma	T2N0M0	2	IA	H7	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
C7	41	F	Invasive ductal carcinoma	T2N0M0	2	IA	H8	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
C8	40	F	Invasive ductal carcinoma	T2N0M0	2	IA	H9	45	F	Invasive ductal carcinoma	T2N0M0	2	IA
C9	40	F	Invasive ductal carcinoma	T2N0M0	2	IA	H10	51	F	Invasive ductal carcinoma	T2N0M0	2	IA
C10	42	F	Invasive ductal carcinoma	T2N1M0	2	IB	H11	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
C11	57	F	Invasive ductal carcinoma	T2N0M0	2	IA	I1	45	F	Invasive ductal carcinoma	T2N1M0	2	IB
D1	49	F	Invasive ductal carcinoma	T2N0M0	2	IA	I2	66	F	Invasive ductal carcinoma	T2N0M0	2	IA
D2	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	I3	75	F	Invasive ductal carcinoma	T2N1M0	2	IB
D3	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	I4	58	F	Invasive ductal carcinoma	T2N1M0	2	IB
D4	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	I5	54	F	Invasive ductal carcinoma	T2N0M0	2	IA
D5	52	F	Invasive ductal carcinoma	T2N1M0	2	IB	I6	54	F	Invasive ductal carcinoma	T2N1M0	2	IB
D6	52	F	Invasive ductal carcinoma	T2N1M0	2	IB	I7	52	F	Invasive ductal carcinoma	T2N1M0	2	IB
D7	52	F	Invasive ductal carcinoma	T2N0M0	2	IA	I8	52	F	Invasive ductal carcinoma	T2N0M0	2	IA
D8	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	I9	59	F	Invasive ductal carcinoma	T2N0M0	2	IA
D9	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	I10	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
D10	44	F	Invasive ductal carcinoma	T2N1M0	2	IB	I11	57	F	Invasive ductal carcinoma	T2N0M0	2	IA
E1	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J1	73	F	Invasive ductal carcinoma	T2N0M0	2	IA
E2	57	F	Invasive ductal carcinoma	T2N0M0	2	IA	J2	66	F	Invasive ductal carcinoma	T2N0M0	2	IA
E3	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	J3	45	F	Invasive ductal carcinoma	T2N0M0	2	IA
E4	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J4	44	F	Invasive ductal carcinoma	T2N0M0	2	IA
E5	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J5	48	F	Invasive ductal carcinoma	T2N0M0	2	IA
E6	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J6	52	F	Invasive ductal carcinoma	T2N0M0	2	IA
E7	58	F	Invasive ductal carcinoma	T2N0M0	2	IA	J7	56	F	Invasive ductal carcinoma	T2N0M0	2	IA
E8	58	F	Invasive ductal carcinoma	T2N0M0	2	IA	J8	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
E9	58	F	Invasive ductal carcinoma	T2N1M0	2	IB	J9	41	F	Invasive ductal carcinoma	T2N1M0	2	IB
E10	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	J10	53	F	Invasive ductal carcinoma	T2N1M0	2	IB
E11	52	F	Invasive ductal carcinoma	T2N1M0	2	IB	J11	57	F	Invasive ductal carcinoma	T2N0M0	2	IA
						K1	45	F	Invasive ductal carcinoma	T2N0M0	2	IA	
						K2	45	F	Invasive ductal carcinoma	T2N0M0	2	IA	
						K3	45	F	Invasive ductal carcinoma	T2N1M0	2	IB	
						K4	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	

Fig. 115B

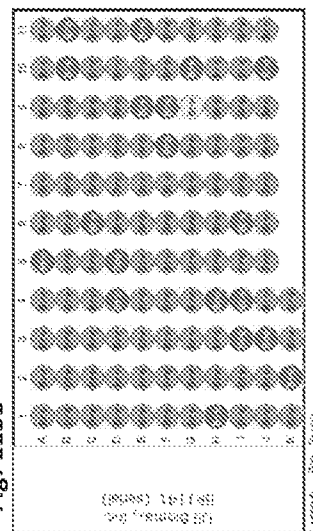
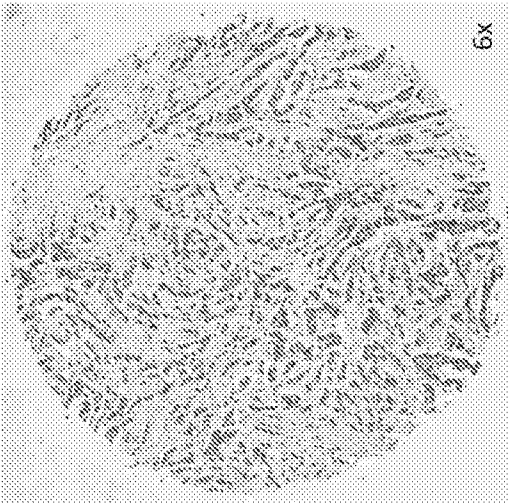


Figure 115A-115C

**PSM/GFR-antibody 20A10 IgG 0.25 ug/mL**  
Breast cancer tissue array BR1141

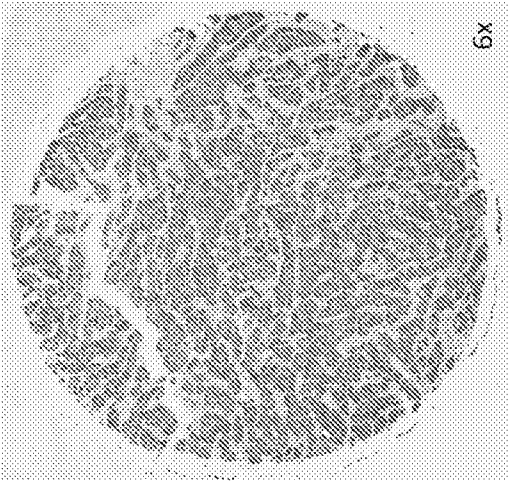
**Fig. 116A**

Position: A4  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



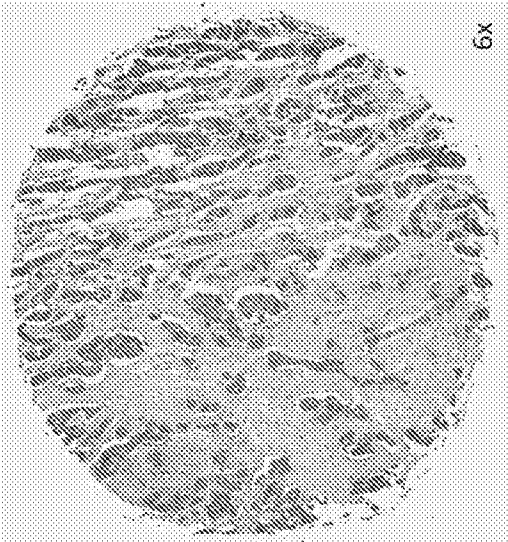
**Fig. 116B**

Position: A11  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0

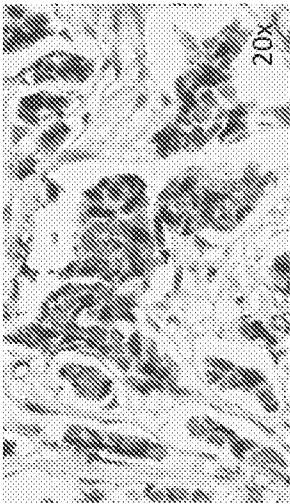


**Fig. 116C**

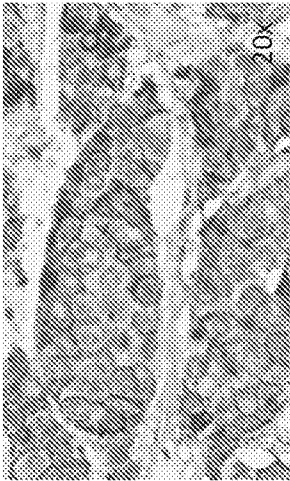
Position: G11  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



**Fig. 116D**



**Fig. 116E**



**Fig. 116F**



**Fig. 116A -- 116F**

Figure 1 displays a 10x10 grid of 100 circular images, labeled A through T on the left and 1 through 10 on the top. Each image shows a different pattern or texture, likely generated by a neural network. The patterns are diverse, ranging from abstract shapes to more recognizable forms like faces and objects.

[illegible]

3665554 - 3665557

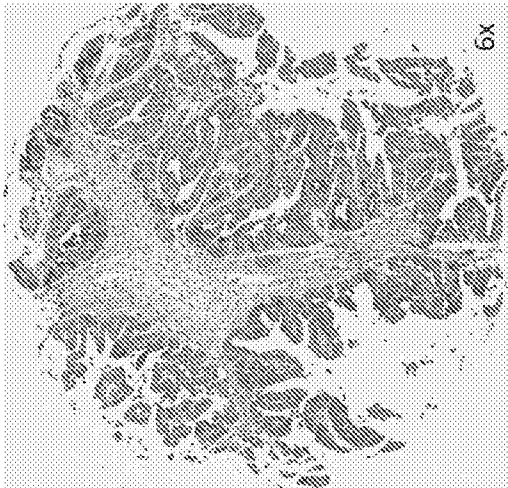
[illegible][illegible]

Figure 117A-117C

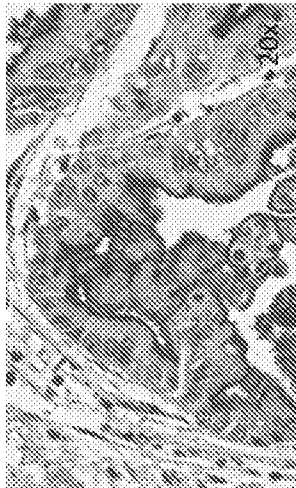
**PSMGR-antibody 20A10 IgG 0.25 ug/mL**  
Pancreatic cancer tissue array PA805c

**Fig. 118A**

Position: D7  
Cell Type: Papillary Adenocarcinoma  
Tumor Grade: 2  
TNM: T2N0M0

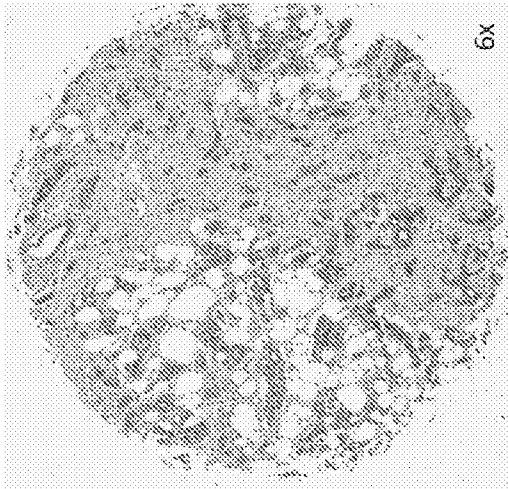


**Fig. 118D**



**Fig. 118B**

Position: E5  
Cell Type: Ductal Carcinoma  
Tumor Grade: 2-3  
TNM: T3N1M0

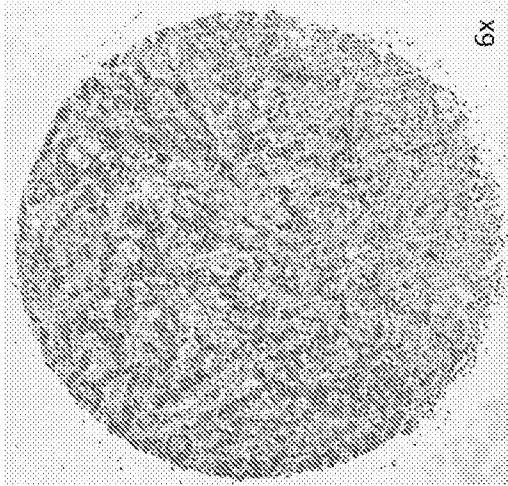


**Fig. 118E**

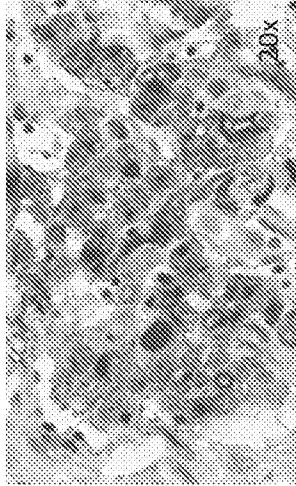


**Fig. 118C**

Position: F1  
Cell Type: Adenocarcinoma  
Tumor Grade: 3  
TNM: T3N0M0



**Fig. 118F**



**Fig. 118A-118F**

PSM/GFR-antibody 20A10 IgG 0.25 ug/mL  
Esophageal cancer tissue array BC001113a

Fig. 119A

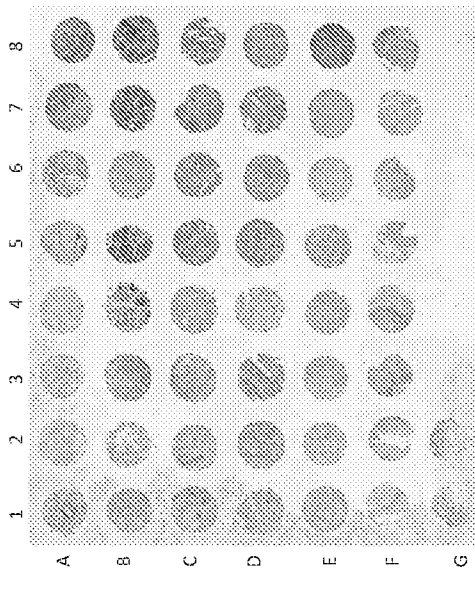


Fig. 119C

Primary Site	Age	Sex	Diagnosis	Primary	Stages	Site	Diagnosis	Primary	Stages	Site	Diagnosis	Primary	Stages	Site	Diagnosis
A1	72	M	Esophagus	Adenocarcinoma	T1NM0	2	Malignant	D1	55	M	Cardia	Adenocarcinoma	T1NM0	2	Malignant
A2	33	F	Esophagus	Adenocarcinoma	T1NM0	2-3	Malignant	D2	70	M	Cardia	Adenocarcinoma	T1NM0	2	Malignant
A3	61	M	Esophagus	Adenocarcinoma	T1NM0	2	Malignant	D3	50	M	Cardia	Adenocarcinoma	T1NM0	2	Malignant
A4	71	M	Esophagus	Adenocarcinoma	T1NM0	2-3	Malignant	D4	64	M	Cardia	Adenocarcinoma	T1NM0	2	Malignant
A5	68	M	Esophagus	Adenocarcinoma	T1NM0	3	Malignant	D5	54	F	Cardia	Adenocarcinoma	T1NM0	3	Malignant
A6	58	F	Esophagus	Adenocarcinoma	T1NM0	3	Malignant	D6	59	M	Cardia	Adenocarcinoma	T1NM0	3	Malignant
A7	51	F	Esophagus	Adenocarcinoma	T1NM0	3	Malignant	D7	72	M	Cardia	Adenocarcinoma	T1NM0	2	Malignant
A8	65	M	Esophagus	Adenocarcinoma	T1NM0	3	Malignant	D8	60	M	Cardia	Adenocarcinoma	T1NM0	3	Malignant
B1	63	M	Esophagus	Adenocarcinoma	T1NM0	3	Malignant	E1	64	M	Cardia	Adenocarcinoma	T1NM0	3	Malignant
B2	61	M	Esophagus	Adenocarcinoma	T1NM0	3	Malignant	E2	42	M	Cardia	Adenocarcinoma	T1NM0	3	Malignant
B3	36	M	Cardia	Adenocarcinoma	T1NM0	1	Malignant	E3	73	M	Cardia	Adenocarcinoma	T1NM0	3	Malignant
B4	55	F	Cardia	Adenocarcinoma	T1NM0	1	Malignant	E4	41	M	Cardia	Adenocarcinoma	T1NM0	3	Malignant
B5	67	M	Cardia	Adenocarcinoma	T1NM0	1	Malignant	E5	63	F	Cardia	Adenocarcinoma	T1NM0	3	Malignant
B6	69	F	Cardia	Adenocarcinoma	T1NM0	1	Malignant	E6	64	F	Cardia	Adenocarcinoma	T1NM0	3	Malignant
B7	69	M	Cardia	Adenocarcinoma	T1NM0	1	Malignant	E7	37	F	Cardia	Adenocarcinoma	T1NM0	3	Malignant
B8	43	M	Cardia	Adenocarcinoma	T1NM0	1	Malignant	E8	51	F	Cardia	Adenocarcinoma	T1NM0	3	Malignant
C1	63	F	Cardia	Adenocarcinoma	T1NM0	1	Malignant	F1	64	F	Esophagus	Adjacent normal	-	-	NAT
C2	54	F	Cardia	Adenocarcinoma	T1NM0	1-2	Malignant	F2	59	M	Esophagus	Adjacent normal	-	-	NAT
C3	30	F	Cardia	Adenocarcinoma	T1NM0	3	Malignant	F3	67	M	Esophagus	Adjacent normal	-	-	NAT
C4	61	M	Cardia	Adenocarcinoma	T1NM0	2	Malignant	F4	43	M	Esophagus	Cancer adjacent	-	-	AT
C5	62	M	Cardia	Adenocarcinoma	T1NM0	2	Malignant	F5	72	M	Esophagus	Cancer adjacent	-	-	AT
C6	65	M	Cardia	Adenocarcinoma	T1NM0	3	Malignant	F6	53	M	Cardia	Adjacent normal, chronic inflammation	-	-	inflammation
C7	72	M	Cardia	Adenocarcinoma	T1NM0	3	Malignant	F7	61	M	Cardia	Adjacent normal, chronic inflammation	-	-	inflammation
C8	74	M	Cardia	Adenocarcinoma	T1NM0	2	Malignant	F8	62	M	Cardia	Adjacent normal, chronic inflammation	-	-	inflammation
							G1	58	M	Cardia	Adjacent normal	-	-	NAT	
							G2	66	M	Cardia	Adjacent normal, chronic inflammation	-	-	inflammation	

Fig. 119B

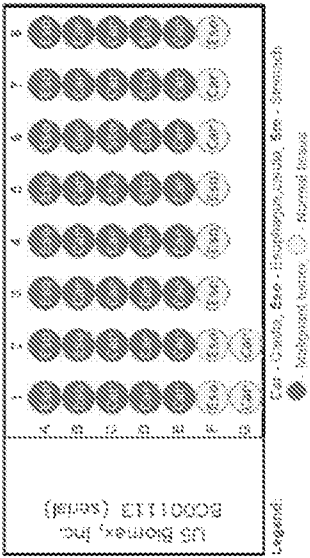


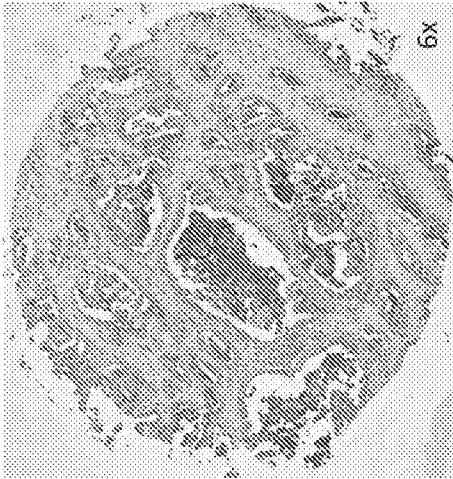
Figure 119A-119C



**PSMGFR-antibody 20A10 IgG 0.25 ug/mL**  
Esophageal cancer tissue array BC001113a

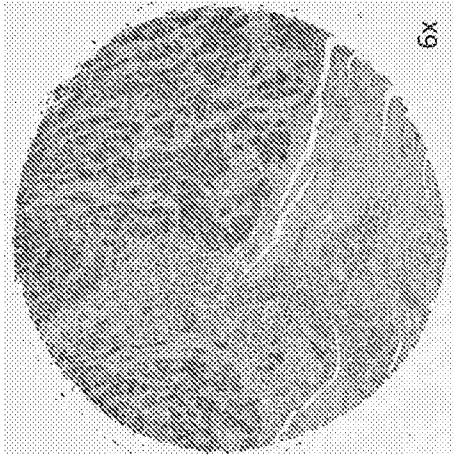
**Fig. 120A**

Position: A1  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N1M0



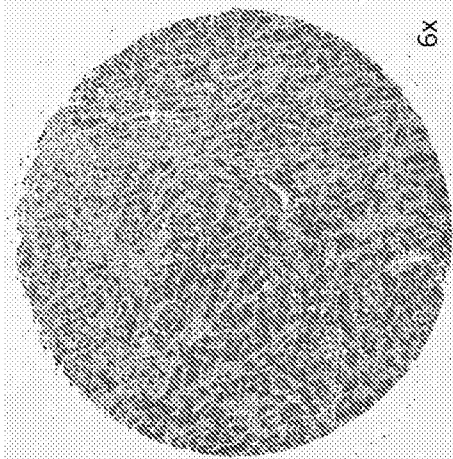
**Fig. 120B**

Position: A7  
Cell Type: Adenocarcinoma  
Tumor Grade: 3  
TNM: T3N0M0

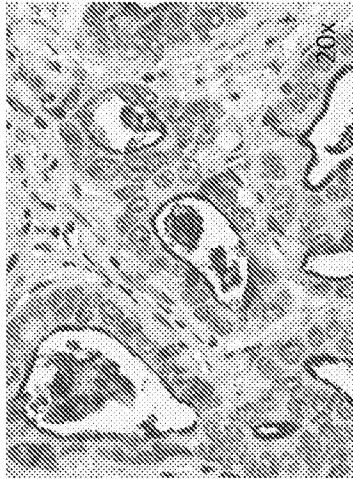


**Fig. 120C**

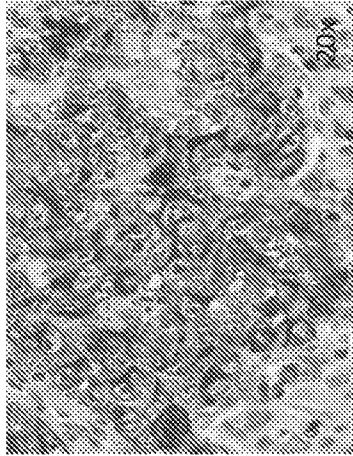
Position: A8  
Cell Type: Adenocarcinoma  
Tumor Grade: 3  
TNM: T4N1M0



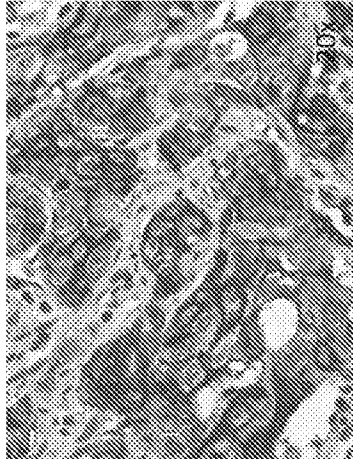
**Fig. 120D**



**Fig. 120E**



**Fig. 120F**



**Fig. 120A – 120F**

PSM/GFR-antibody 3C2B1 20 ug/mL  
FDA Normal tissue array MNO1021

Fig. 121A

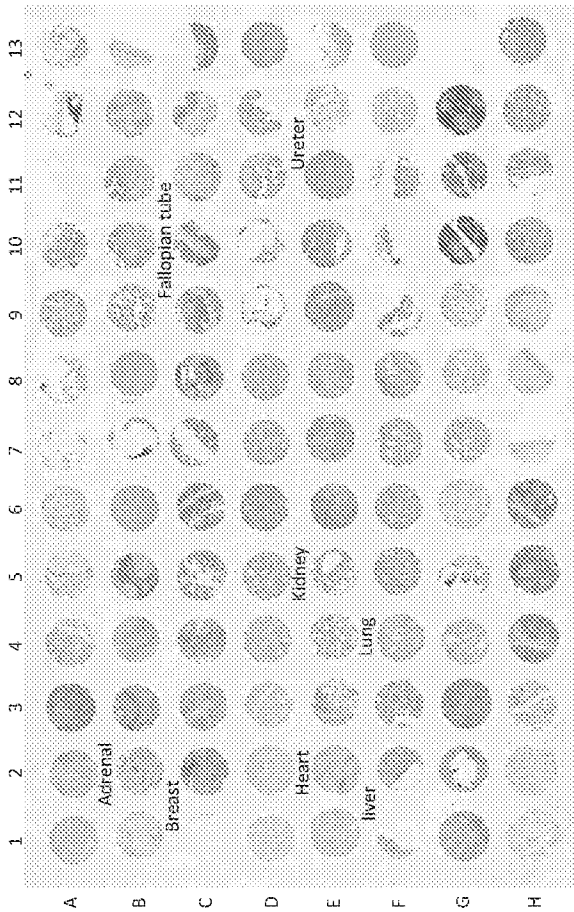


Fig. 121C

Position	Age	Sex	Organ/Anatomic Site	Pathology diagnosis	Organ/Anatomic Site	Position	Age	Sex	Pathology diagnosis
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	53	Adrenal gland	Normal	E3	M	72	Lung	Normal
A4	M	75	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	43	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bone, bone marrow	Normal	E7	F	46	Ovary	Normal
A8	M	43	Bone, bone marrow	Normal	E8	F	40	Ovary	Normal
A9	F	44	Head & neck, salivary gland	Normal	E9	M	42	Pancreas	Normal
A10	M	18	Head & neck, salivary gland	Normal	E10	F	28	Pancreas	Normal
A11	M	50	Eye	Normal	E11	F	59	Pancreas	Normal
A12	M	50	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	F	35	Breast	Normal	E13	M	72	Parathyroid	Adenoma
B1	F	33	Breast	Normal	F1	F	3	Pituitary gland	Normal
B2	F	33	Breast	Normal	F2	F	32	Pituitary gland	Normal
B3	M	58	Brain, cerebellum	Normal	F3	F	30	Pituitary gland	Normal
B4	M	95	Brain, cerebellum	Normal	F4	F	27	Pituitary gland	Normal
B5	F	58	Brain, cerebellum	Normal	F5	F	27	Pituitary gland	Normal
B6	M	58	Brain, cerebral cortex	Normal	F6	M	64	Prostate	Normal
B7	M	85	Brain, cerebral cortex	Normal	F7	M	65	Prostate	Normal
B8	F	58	Brain, cerebral cortex	Normal	F8	M	65	Prostate	Normal
B9	F	42	Fallopian tube	Normal	F9	F	32	Skin	Normal
B10	F	32	Fallopian tube	Normal	F10	F	66	Skin	Normal
B11	F	24	Fallopian tube	Normal	F11	M	28	Skin	Normal
B12	F	45	Esophagus	Normal	F12	M	47	Spinal cord	Normal
B13	F	68	Esophagus	Normal	F13	M	58	Spinal cord	Normal
C1	M	54	Esophagus	Normal	G1	M	27	Spleen	Normal
C2	M	45	Stomach	Normal	G2	M	60	Spleen	Normal
C3	M	49	Stomach	Normal	G3	M	31	Spleen	Normal
C4	M	77	Stomach	Normal	G4	F	60	Skeletal muscle	Normal
C5	M	45	Intestine, small intestine	Normal	G5	F	49	Skeletal muscle	Normal
C6	F	75	Intestine, small intestine	Normal	G6	M	70	Skeletal muscle	Normal
C7	F	2	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C8	M	2	Intestine, colon	Normal	G8	M	43	Testis	Normal
C9	M	48	Intestine, colon	Normal	G9	M	77	Testis	Normal
C10	M	74	Intestine, rectum	Normal	G10	M	15	Thymus	Normal
C11	F	47	Intestine, rectum	Normal	G11	M	28	Thymus	Normal
C12	M	86	Intestine, rectum	Normal	G12	F	9	Thymus	Normal
C13	M	58	Heart	Normal	G13	-	-	-	-
D1	M	46	Heart	Normal	H1	F	28	Thyroid	Normal
D2	F	34	Heart	Normal	H2	F	37	Thyroid	Normal
D3	F	52	Kidney, cortex	Normal	H3	F	51	Thyroid	Normal
D4	M	23	Kidney, cortex	Normal	H4	M	46	Tonsil	Normal
D5	M	63	Kidney, cortex	Normal	H5	M	37	Tonsil	Normal
D6	F	52	Kidney, medulla	Normal	H6	M	13	Tonsil	Normal
D7	M	23	Kidney, medulla	Normal	H7	F	38	Uterus, cervix	Normal
D8	M	59	Peripheral nerve	Schwannoma	H8	F	35	Uterus, cervix	Normal
D9	F	28	Peripheral nerve	Schwannoma	H9	F	44	Uterus, cervix	Normal
D10	F	44	Ureter	Normal	H10	F	36	Uterus, endometrium	Normal
D11	F	42	Ureter	Normal	H11	F	41	Uterus, endometrium	Normal
D12	F	57	Liver	Normal	H12	F	46	Uterus, endometrium	Normal
D13	F	57	Liver	Normal	H13	M	58	Skin	Malignant melanoma (tissue marker)

Fig. 121B

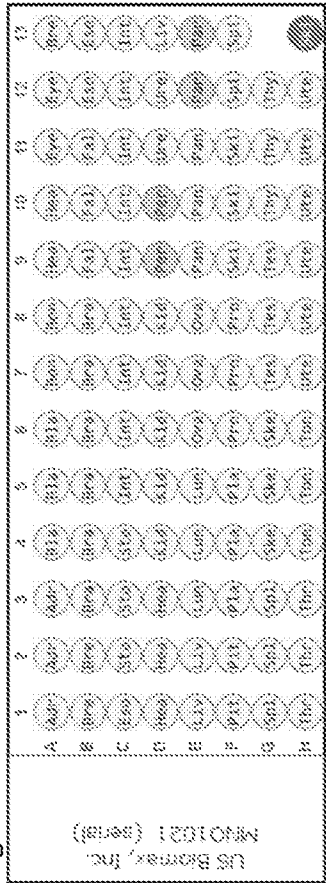
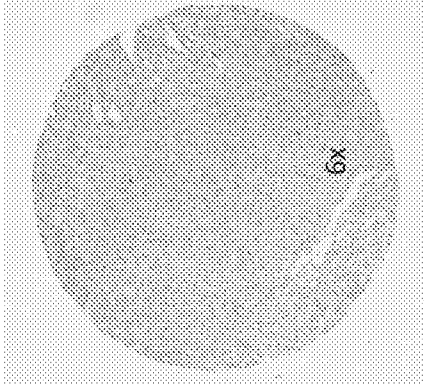


Figure 121A-121C

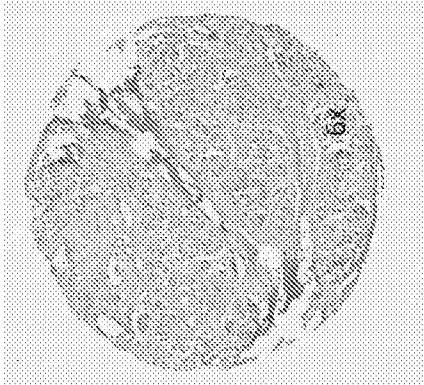


**PSMGFR-antibody 3C2B1 IgG 20 ug/mL**  
**FDA Normal tissue array MNO1021**

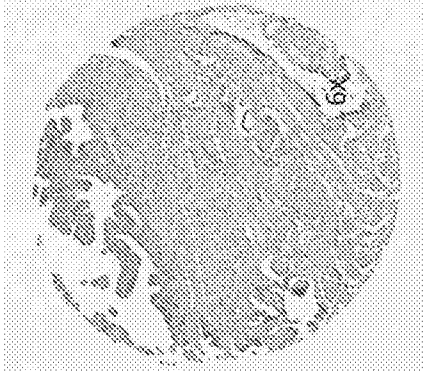
**Fig. 122A**  
**Normal Adrenal Gland**  
**A1**



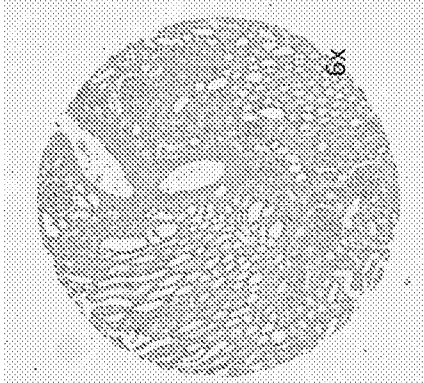
**Fig. 122B**  
**Normal Breast**  
**B2**



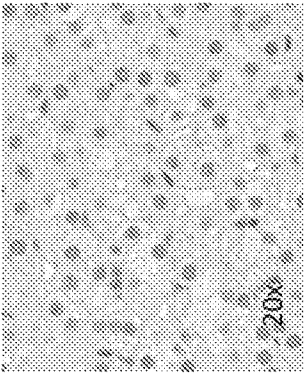
**Fig. 122C**  
**Normal Fallopian Tubes**  
**B10**



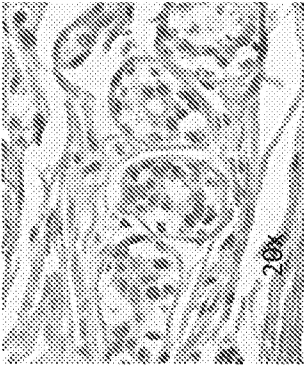
**Fig. 122D**  
**Normal Kidney**  
**D5**



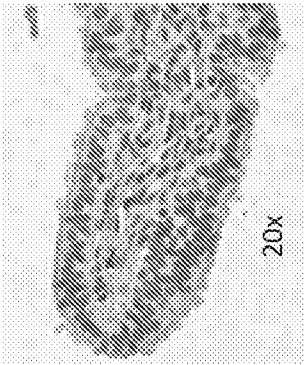
**Fig. 122E**



**Fig. 122F**



**Fig. 122G**



**Fig. 122H**



**Figure 122A-122H**

PSMGR-antibody 3C2B1 IgG 20 ug/mL  
FDA Normal tissue array MNO1021

Fig. 122I

Normal Heart

D3

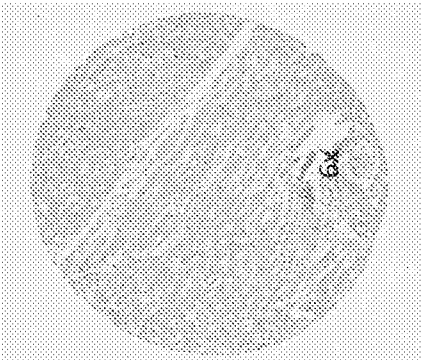


Fig. 122J

Normal Liver

D13

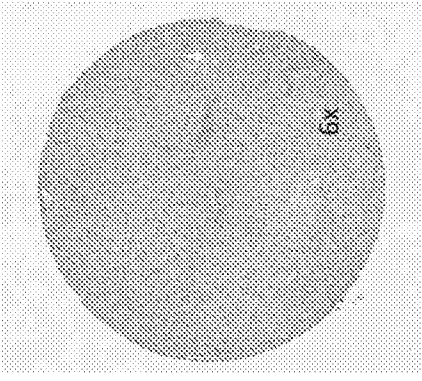


Fig. 122K

Normal Lung

E3

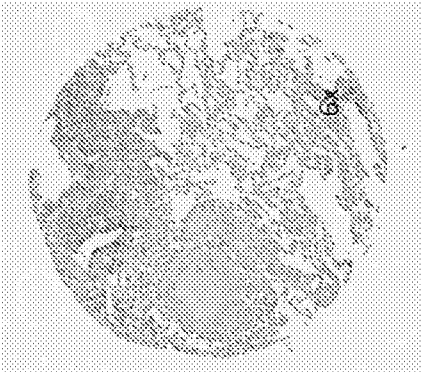


Fig. 122L

Normal Ureter

D11



Fig. 122M

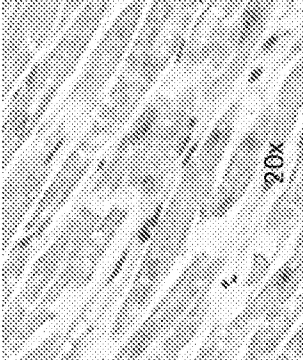


Fig. 122N

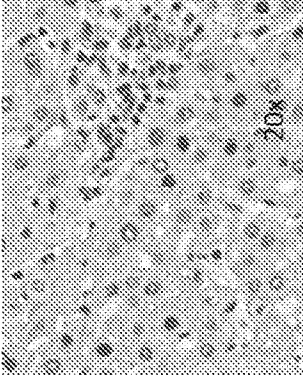


Fig. 122O

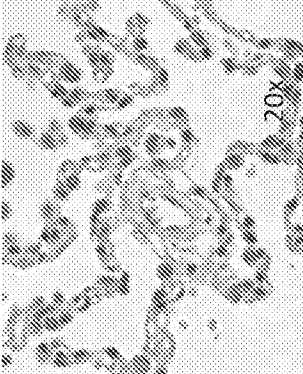


Fig. 122P

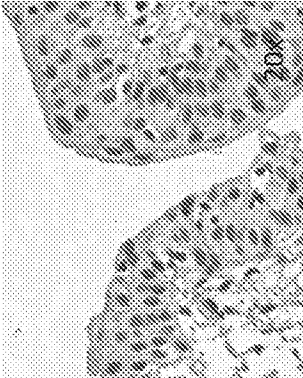


Figure 122I-122P

**PSMGFR-antibody 3C2B1 IgG 20 ug/mL**  
**FDA Normal tissue array MNO1021**

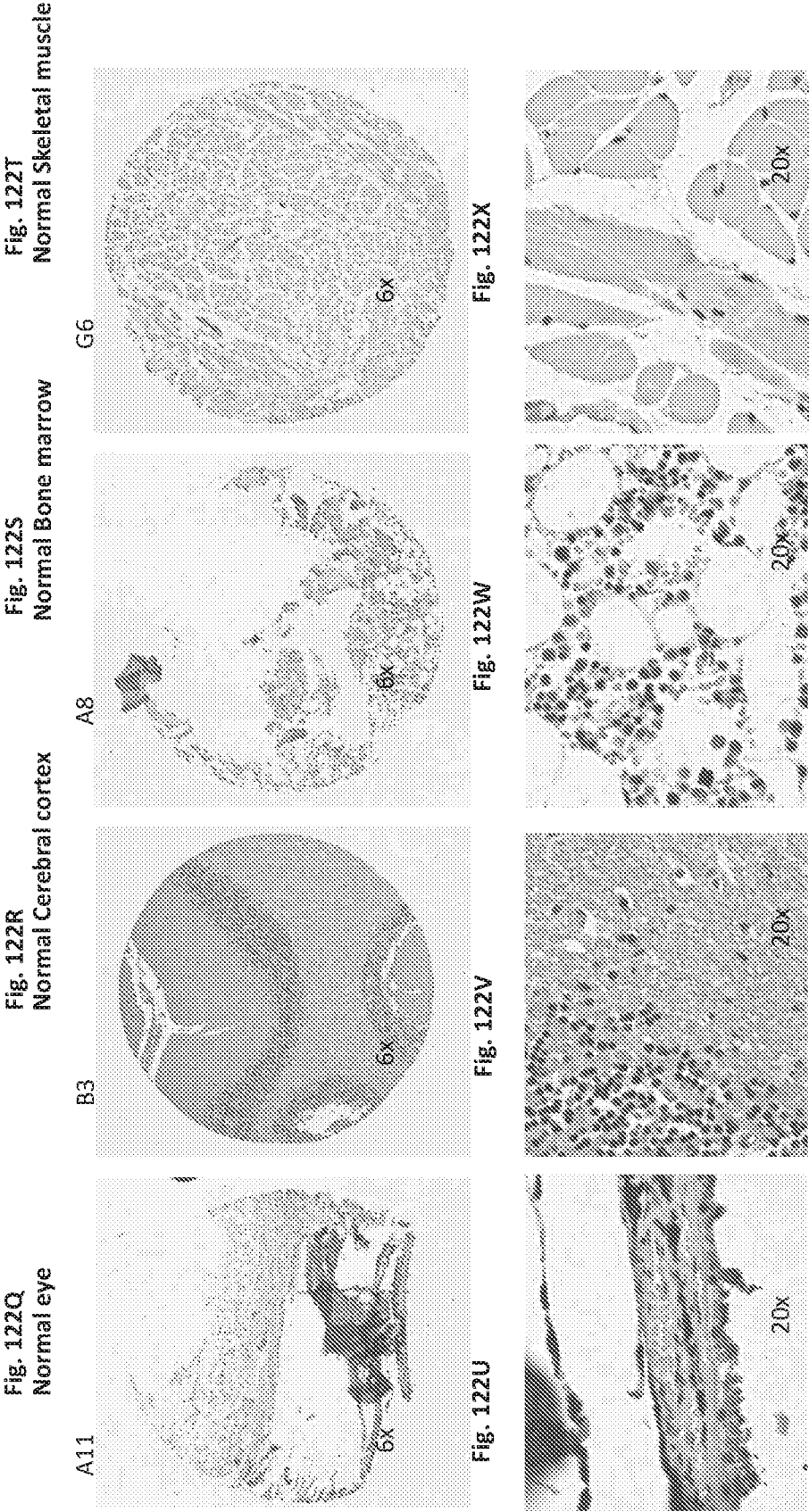


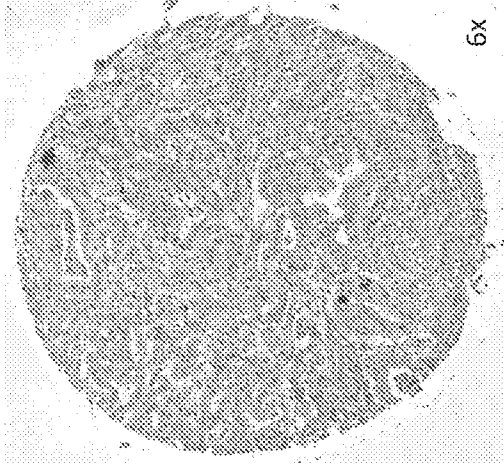
Figure 122Q-122X



**PSMGFR-antibody 3C2B1 IgG 20 ug/mL**  
**Pancreatic cancer tissue array PA1003**

**Fig. 124A**

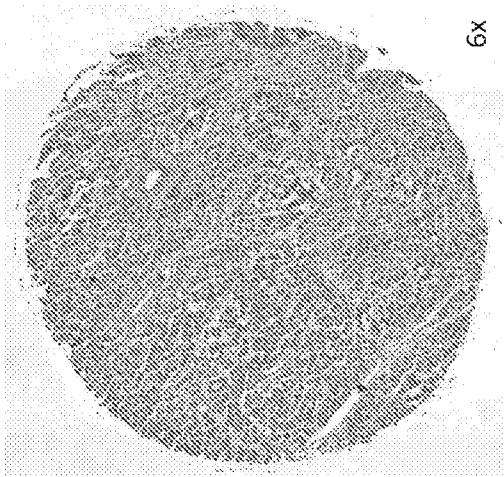
Position: A2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



6x

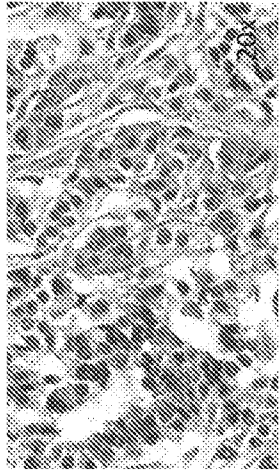
**Fig. 124B**

Position: B2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



6x

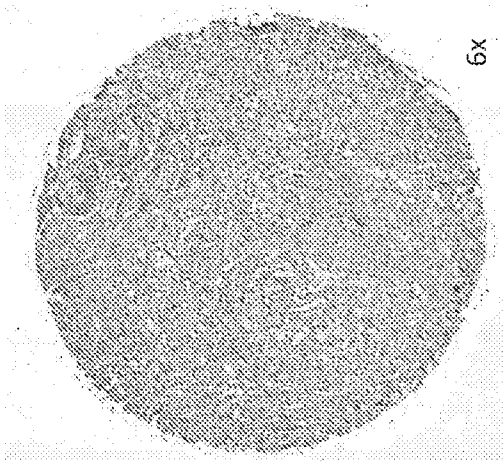
**Fig. 124E**



20x

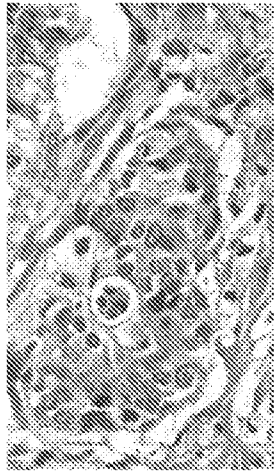
**Fig. 124C**

Position: C2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



6x

**Fig. 124F**



20x

**Fig. 124D**



20x

**Fig. 124D-124F**

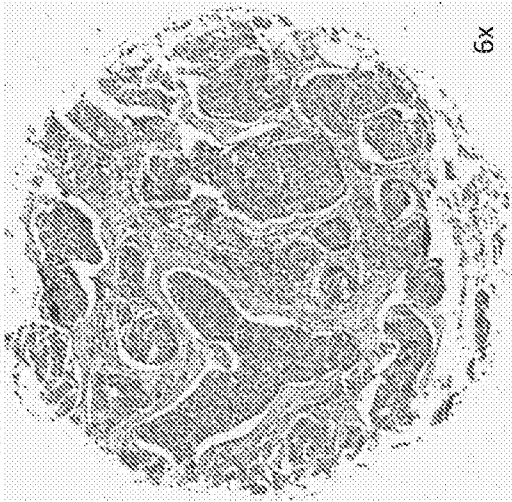




**PSMGFR-antibody 3C2B1 20 ug/mL**  
**Breast cancer tissue array BR1141a**

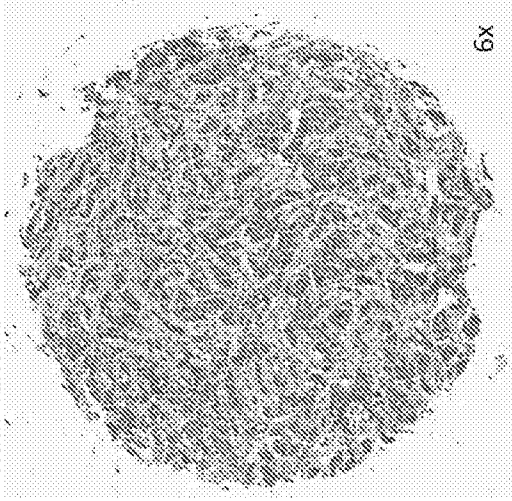
**Fig. 126A**

Position: C6  
Cell Type: Invasive carcinoma  
Tumor Grade: 2  
TNM: T3N2M0



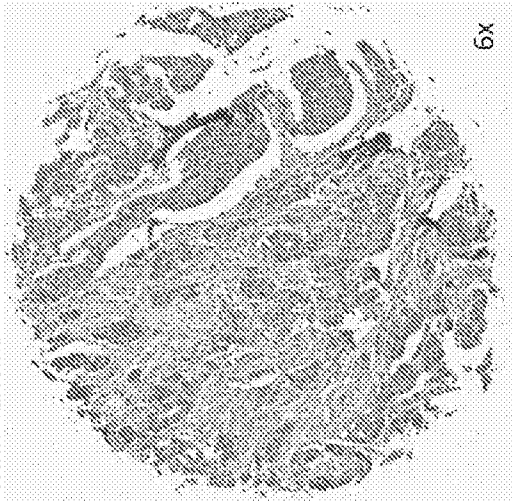
**Fig. 126B**

Position: D6  
Cell Type: Invasive carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



**Fig. 126C**

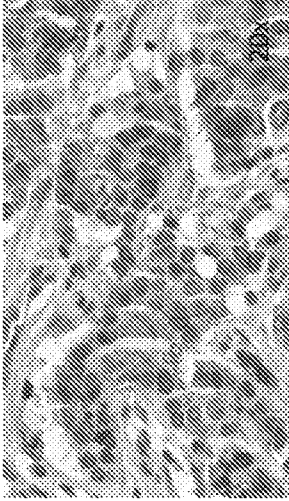
Position: E1  
Cell Type: Invasive carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



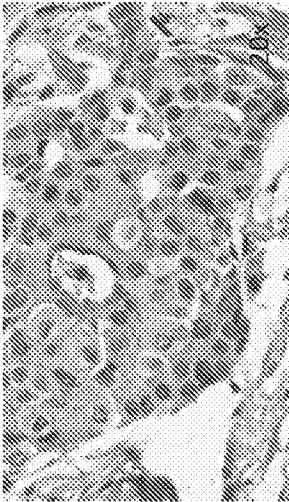
**Fig. 126D**



**Fig. 126E**



**Fig. 126F**



**Fig. 126A-126F**

PSMGFR-antibody 5C6F3 1 ug/mL  
FDA Normal tissue array MNO1021

Fig. 127A

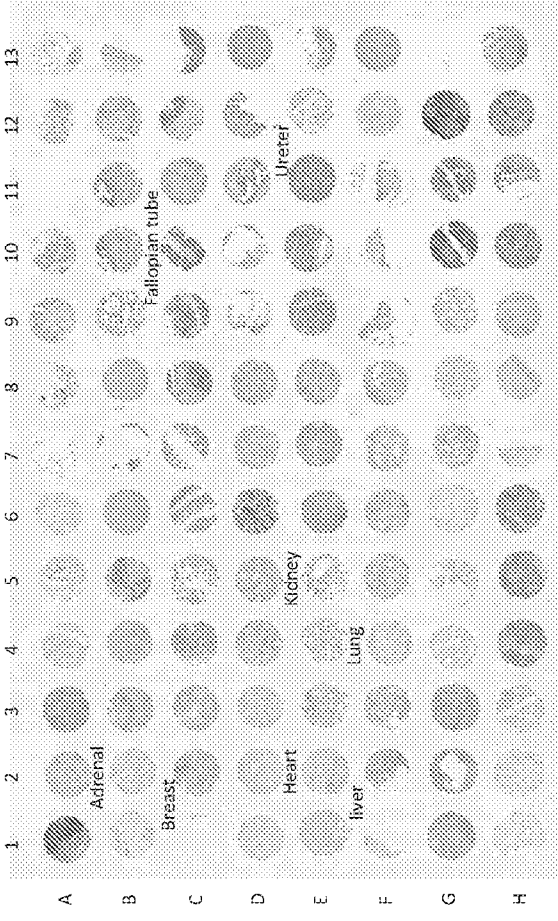


Fig. 127C

Position	Age	Sex	Organ/Anatomic Site	Pathology diagnosis	Position	Age	Sex	Organ/Anatomic Site	Pathology diagnosis
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	53	Adrenal gland	Normal	E3	M	72	Liver	Normal
A4	M	73	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	52	Bladder, urinary	Normal	E5	M	43	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bladder, urinary	Normal	E7	F	48	Ovary	Normal
A8	M	43	Bone, bone marrow	Normal	E8	F	40	Ovary	Normal
A9	M	44	Bone, bone marrow	Normal	E9	M	42	Pancreas	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	35	Pituitary gland	Normal
A11	M	18	Head and neck, salivary gland	Normal	E11	F	35	Pituitary gland	Normal
A12	M	55	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	M	35	Breast	Normal	E13	M	72	Parathyroid	Adenoma
A14	F	38	Breast	Normal	F1	F	3	Pituitary gland	Normal
A15	F	38	Breast	Normal	F2	F	3	Pituitary gland	Normal
A16	M	35	Brain, cerebellum	Normal	F3	F	32	Placenta	Normal
A17	M	35	Brain, cerebellum	Normal	F4	F	30	Placenta	Normal
A18	M	35	Brain, cerebellum	Normal	F5	F	30	Placenta	Normal
A19	M	35	Brain, cerebellum	Normal	F6	M	64	Prostate	Normal
A20	M	35	Brain, cerebellum	Normal	F7	M	64	Prostate	Normal
A21	F	42	Fallopian tube	Normal	F8	M	64	Prostate	Normal
A22	F	42	Fallopian tube	Normal	F9	M	64	Prostate	Normal
A23	F	42	Fallopian tube	Normal	F10	M	64	Prostate	Normal
A24	F	42	Fallopian tube	Normal	F11	M	64	Prostate	Normal
A25	F	42	Fallopian tube	Normal	F12	M	64	Prostate	Normal
A26	F	42	Fallopian tube	Normal	F13	M	64	Prostate	Normal
A27	F	42	Fallopian tube	Normal	G1	M	27	Spleen	Normal
A28	F	42	Fallopian tube	Normal	G2	M	27	Spleen	Normal
A29	F	42	Fallopian tube	Normal	G3	M	27	Spleen	Normal
A30	F	42	Fallopian tube	Normal	G4	M	27	Spleen	Normal
A31	M	49	Stomach	Normal	G5	M	27	Spleen	Normal
A32	M	49	Stomach	Normal	G6	M	27	Spleen	Normal
A33	M	49	Stomach	Normal	G7	M	27	Spleen	Normal
A34	M	49	Stomach	Normal	G8	M	27	Spleen	Normal
A35	M	49	Stomach	Normal	G9	M	27	Spleen	Normal
A36	M	49	Stomach	Normal	G10	M	27	Spleen	Normal
A37	M	49	Stomach	Normal	G11	M	27	Spleen	Normal
A38	M	49	Stomach	Normal	G12	M	27	Spleen	Normal
A39	M	49	Stomach	Normal	G13	M	27	Spleen	Normal
A40	M	49	Stomach	Normal	H1	F	26	Thyroid	Normal
A41	M	49	Stomach	Normal	H2	F	26	Thyroid	Normal
A42	M	49	Stomach	Normal	H3	F	26	Thyroid	Normal
A43	M	49	Stomach	Normal	H4	F	26	Thyroid	Normal
A44	M	49	Stomach	Normal	H5	F	26	Thyroid	Normal
A45	M	49	Stomach	Normal	H6	F	26	Thyroid	Normal
A46	M	49	Stomach	Normal	H7	F	26	Thyroid	Normal
A47	M	49	Stomach	Normal	H8	F	26	Thyroid	Normal
A48	M	49	Stomach	Normal	H9	F	26	Thyroid	Normal
A49	M	49	Stomach	Normal	H10	F	26	Thyroid	Normal
A50	M	49	Stomach	Normal	H11	F	26	Thyroid	Normal
A51	M	49	Stomach	Normal	H12	F	26	Thyroid	Normal
A52	M	49	Stomach	Normal	H13	F	26	Thyroid	Normal
A53	M	49	Stomach	Normal	H14	F	26	Thyroid	Normal
A54	M	49	Stomach	Normal	H15	F	26	Thyroid	Normal
A55	M	49	Stomach	Normal	H16	F	26	Thyroid	Normal
A56	M	49	Stomach	Normal	H17	F	26	Thyroid	Normal
A57	M	49	Stomach	Normal	H18	F	26	Thyroid	Normal
A58	M	49	Stomach	Normal	H19	F	26	Thyroid	Normal
A59	M	49	Stomach	Normal	H20	F	26	Thyroid	Normal
A60	M	49	Stomach	Normal	H21	F	26	Thyroid	Normal

Fig. 127B

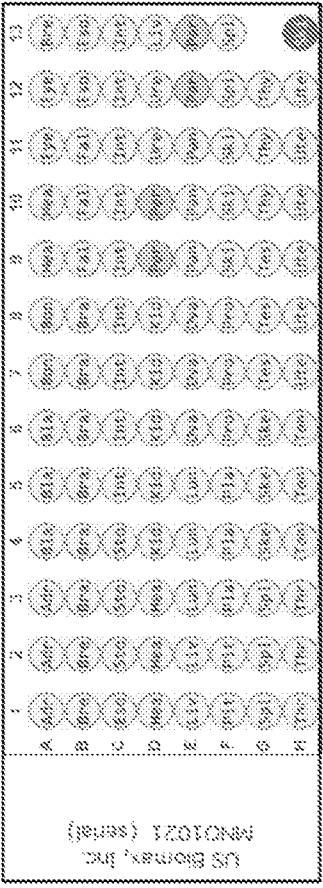


Figure 127A-127C



PSMGFR-antibody 5C6F3 1 ug/mL  
FDA Normal tissue array MNO1021

Fig. 128A

Normal Adrenal Gland

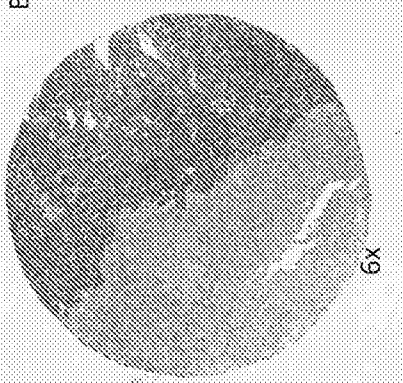


Fig. 128B

Normal Breast

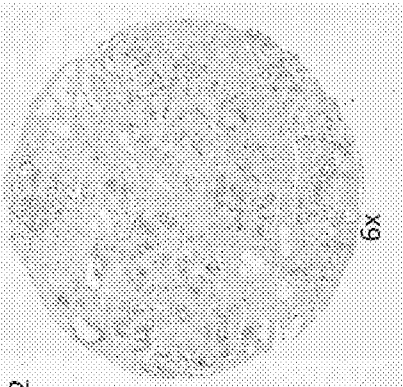


Fig. 128C

Normal Fallopian Tubes

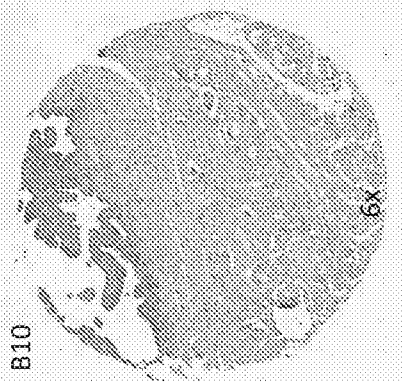


Fig. 128D

Normal Kidney

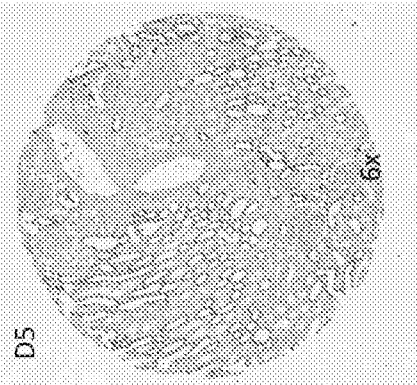


Fig. 128E



Fig. 128F

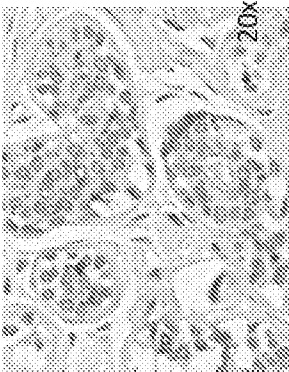


Fig. 128G

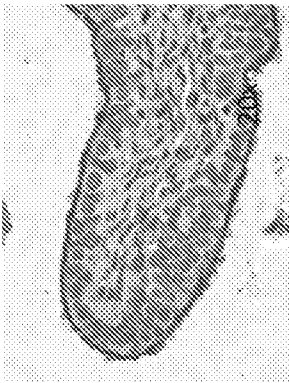


Fig. 128H

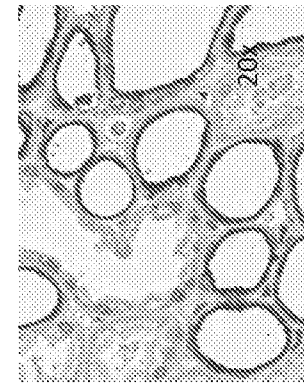


Figure 128A-128H

PSMGFR-antibody 5C6F3 1 ug/mL  
FDA Normal tissue array MNO1021

Fig. 128I  
Normal Heart

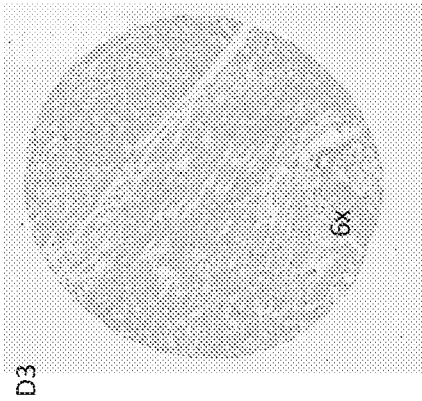


Fig. 128J  
Normal Liver

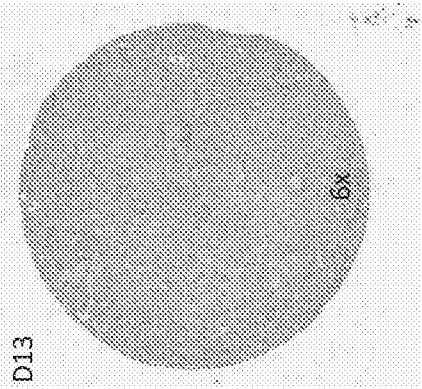


Fig. 128K  
Normal Lung

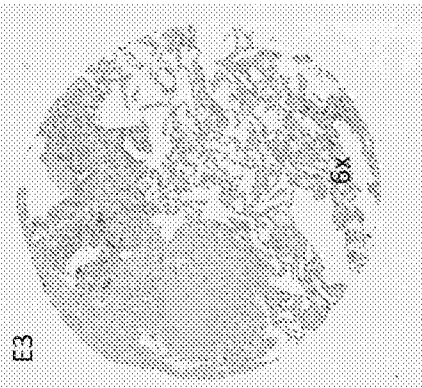


Fig. 128L  
Normal Ureter

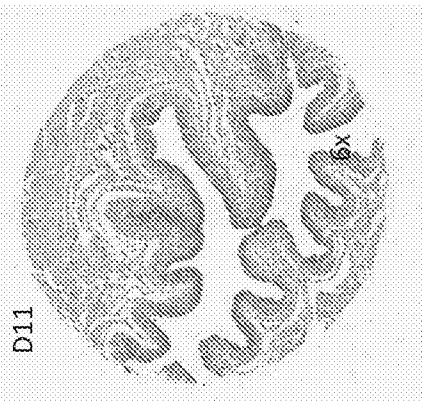


Fig. 128M

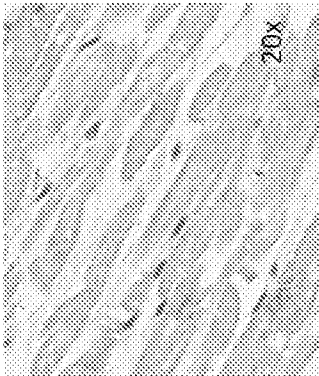


Fig. 128N

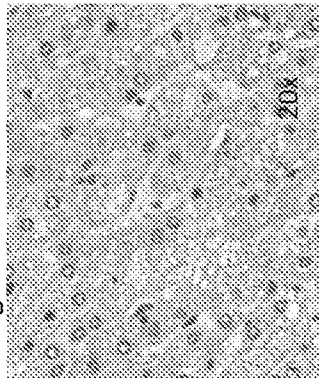


Fig. 128O

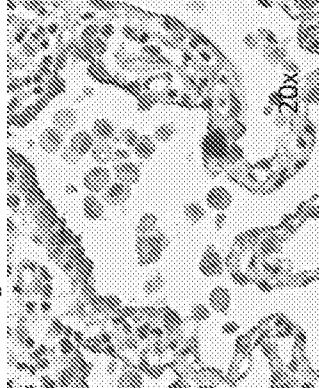


Fig. 128P

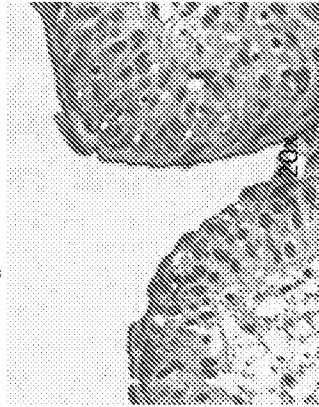


Figure 128I-  
128P

PSMGFR-antibody 5C6F3 1 ug/mL  
FDA Normal tissue array MNO1021

Fig. 128Q  
Normal eye

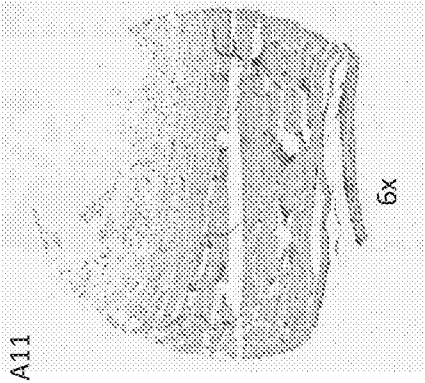


Fig. 128R  
Normal Cerebral cortex

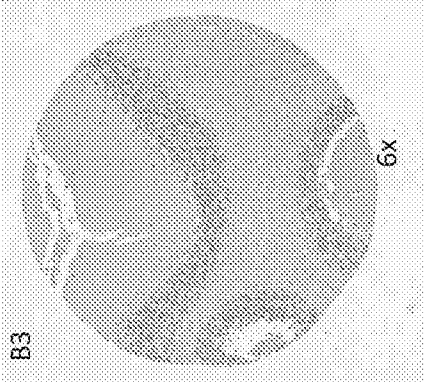


Fig. 128S  
Normal Bone marrow

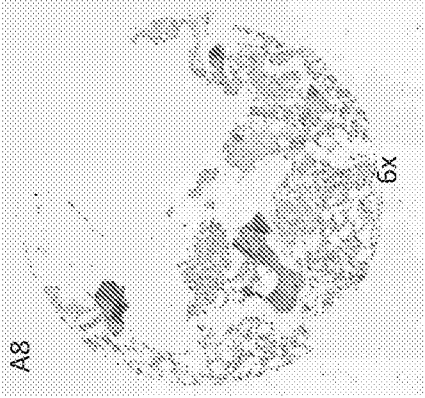


Fig. 128T  
Normal Skeletal muscle

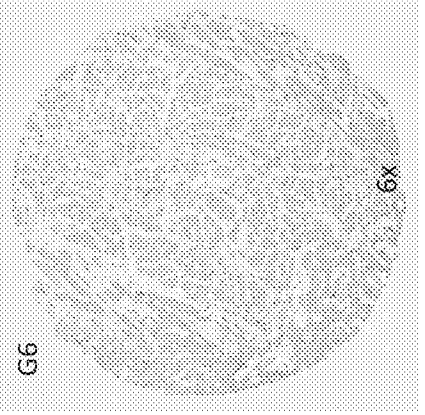


Fig. 128U

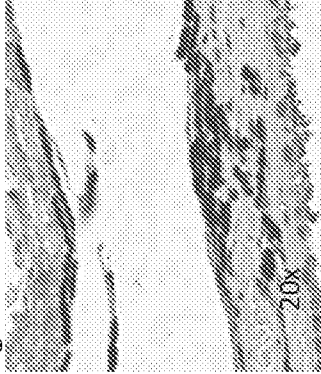


Fig. 128V

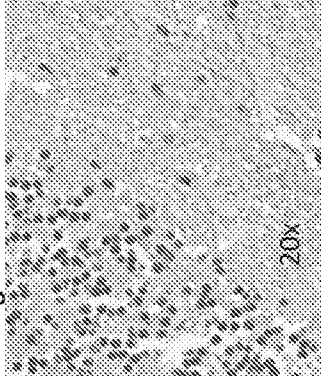


Fig. 128W

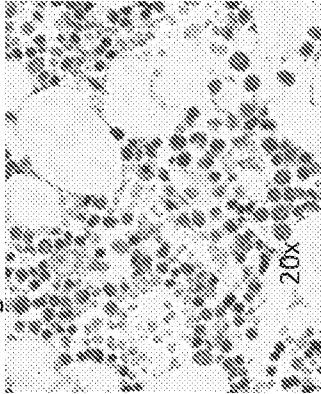


Fig. 128X

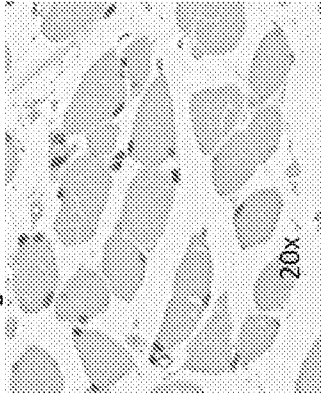


Figure 128Q-128X



**PSMGFR-antibody 5C6F3 1 ug/mL**  
**Pancreatic cancer tissue array PA1003**

**Fig. 130D**

Position: A2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



**Fig. 130D**

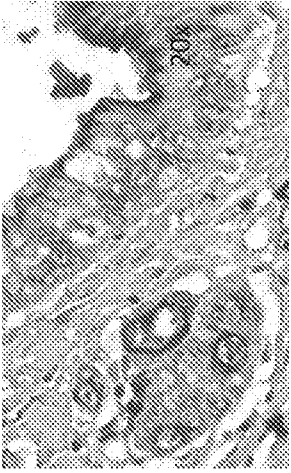


**Fig. 130E**

Position: B2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0

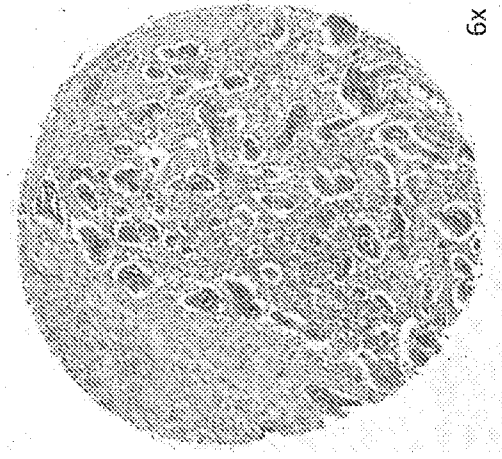


**Fig. 130E**

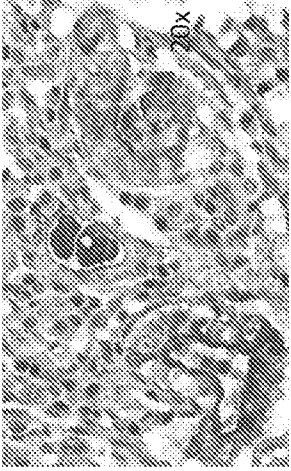


**Fig. 130F**

Position: C2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



**Fig. 130F**



**Fig. 130A-130F**

A 10x10 grid of 100 grayscale images of handwritten digits, labeled 1 through 10 on the left. The digits are generated by a generative model, showing a variety of styles and orientations.

A1	30	I	Investive	cariorhina	2	IA	Malignant	F1	43	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A2	31	I	Investive	cariorhina	2	IA	Malignant	F2	44	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A3	32	I	Investive	cariorhina	2	IA	Malignant	F3	62	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A4	33	I	Investive	cariorhina	2	IB	Malignant	F4	48	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A5	34	I	Investive	cariorhina	2	IB	Malignant	F5	45	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A6	35	I	Investive	cariorhina	2	IB	Malignant	F6	47	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A7	36	I	Investive	cariorhina	2	IA	Malignant	F7	50	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A8	37	I	Investive	cariorhina	2	IB	Malignant	F8	48	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A9	38	I	Investive	cariorhina	2	IB	Malignant	F9	40	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A10	39	I	Investive	cariorhina	2	IA	Malignant	F10	50	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
A11	40	I	Investive	cariorhina	2	IA	Malignant	F11	50	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B1	41	I	Investive	cariorhina	2	IIA	Malignant	G1	47	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B2	42	I	Investive	cariorhina	2	IIA	Malignant	G2	40	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B3	43	I	Investive	cariorhina	2	IA	Malignant	G3	54	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B4	44	I	Investive	cariorhina	2	IA	Malignant	G4	56	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B5	45	I	Investive	cariorhina	2	IA	Malignant	G5	48	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B6	46	I	Investive	cariorhina	2	IB	Malignant	G6	45	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B7	47	I	Investive	cariorhina	2	IB	Malignant	G7	50	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B8	48	I	Investive	cariorhina	2	IA	Malignant	G8	45	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B9	49	I	Investive	cariorhina	2	IB	Malignant	G9	42	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B10	50	I	Investive	cariorhina	2	IA	Malignant	G10	36	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
B11	51	I	Investive	cariorhina	2	IA	Malignant	G11	44	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
C1	52	I	Investive	cariorhina	2	IIA	Malignant	H1	47	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
C2	53	I	Investive	cariorhina	2	IIA	Malignant	H2	50	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
C3	54	I	Investive	cariorhina	2	IA	Malignant	H3	52	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
C4	55	I	Investive	cariorhina	2	IA	Malignant	H4	35	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
C5	56	I	Investive	cariorhina	2	IIA	Malignant	H5	40	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
C6	57	I	Investive	cariorhina	2	IIA	Malignant	H6	46	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
C7	58	I	Investive	cariorhina	2	IB	Malignant	H7	44	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2
C8	59	I	Investive	cariorhina	2	IB	Malignant	H8	45	F	Investive	cariorhina	2	IIA	Malignant	2	IV	Malignant	2

[illegible]

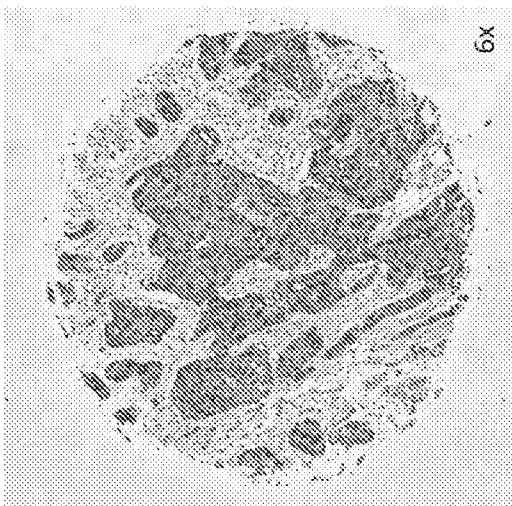
Figure 131A-131C



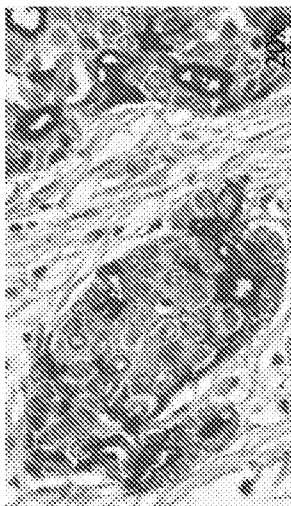
**PSMGFR-antibody 5C6F3 1 ug/mL**  
**Breast cancer tissue array BR1141a**

**Fig. 132A**

Position: B1  
Cell Type: Invasive carcinoma  
Tumor Grade: 2  
TNM: T2N0M0

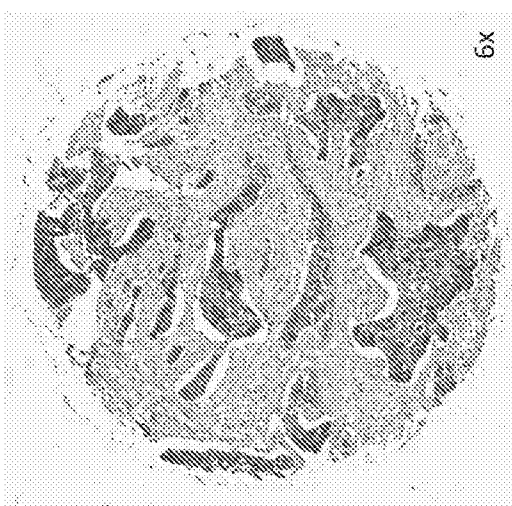


**Fig. 132D**



**Fig. 132B**

Position: B3  
Cell Type: Invasive carcinoma  
Tumor Grade: 2  
TNM: T2N0M0

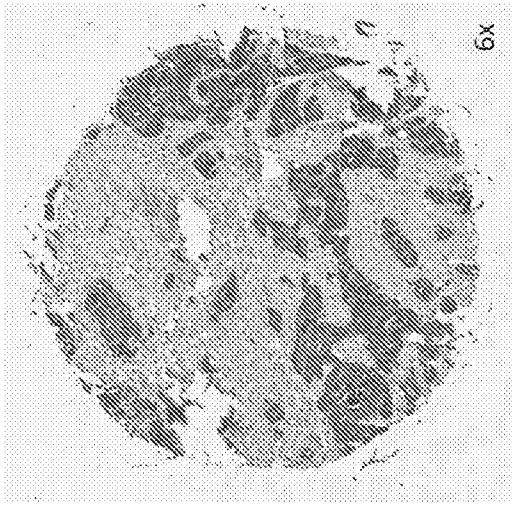


**Fig. 132E**



**Fig. 132C**

Position: B10  
Cell Type: Invasive carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



**Fig. 132F**



**Fig. 132A – 132F**

PSMGFR-antibody 18B4 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 133A

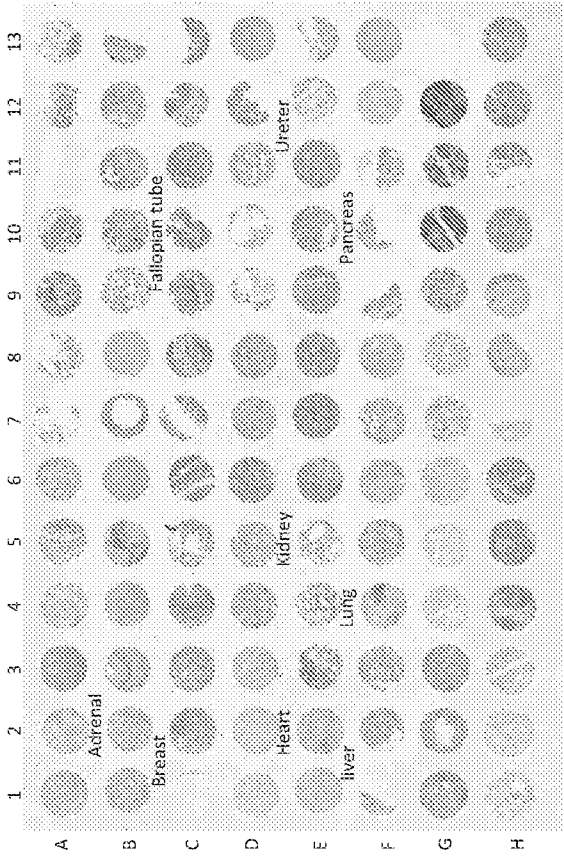


Fig. 133C

Position	Age	Sex	Organ/Anatomic Site	Pathology diagnosis	Position	Age	Sex	Organ/Anatomic Site	Pathology diagnosis
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	53	Adrenal gland	Normal	E3	M	72	Liver	Normal
A4	M	75	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	43	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bone, bone marrow	Normal	E7	F	46	Ovary	Normal
A8	M	43	Bone, bone marrow	Normal	E8	F	40	Ovary	Normal
A9	F	44	Head and neck, salivary gland	Normal	E9	M	42	Pancreas	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	26	Pancreas	Normal
A11	M	55	Eye	Normal	E11	F	59	Pancreas	Normal
A12	M	50	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	F	35	Breast	Normal	E13	M	72	Parathyroid	Adenoma
B1	F	38	Breast	Normal	F1	F	3	Pharynx, pharynx	Normal
B2	F	33	Breast	Normal	F2	F	32	Pharynx, pharynx	Normal
B3	M	58	Brain, cerebrum	Normal	F3	F	30	Placenta	Normal
B4	M	85	Brain, cerebrum	Normal	F4	F	30	Placenta	Normal
B5	F	58	Brain, cerebrum	Normal	F5	F	27	Placenta	Normal
B6	M	58	Brain, cerebral cortex	Normal	F6	M	54	Prostate	Normal
B7	M	85	Brain, cerebral cortex	Normal	F7	M	65	Prostate	Normal
B8	F	42	Fallopian tube	Normal	F8	M	65	Prostate	Normal
B9	F	32	Fallopian tube	Normal	F9	F	32	Skin	Normal
B10	F	24	Fallopian tube	Normal	F10	F	68	Skin	Normal
B11	M	45	Esophagus	Normal	F11	M	28	Skin	Normal
B12	F	88	Esophagus	Normal	F12	M	47	Spinal cord	Normal
B13	M	54	Esophagus	Normal	F13	M	56	Spinal cord	Normal
C1	M	45	Stomach	Normal	G1	M	27	Spleen	Normal
C2	M	49	Stomach	Normal	G2	M	60	Spleen	Normal
C3	M	49	Stomach	Normal	G3	M	31	Spleen	Normal
C4	M	77	Stomach	Normal	G4	F	60	Skeletal muscle	Normal
C5	M	45	Intestine, small intestine	Normal	G5	F	49	Skeletal muscle	Normal
C6	F	75	Intestine, small intestine	Normal	G6	M	70	Skeletal muscle	Normal
C7	F	75	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C8	M	46	Intestine, colon	Normal	G8	M	30	Testis	Normal
C9	M	46	Intestine, colon	Normal	G9	M	77	Testis	Normal
C10	M	74	Intestine, colon	Normal	G10	M	15	Thymus	Normal
C11	F	47	Intestine, rectum	Normal	G11	M	26	Thymus	Normal
C12	F	75	Intestine, rectum	Normal	G12	F	9	Thymus	Normal
C13	M	86	Intestine, rectum	Normal	G13	-	-	-	-
D1	M	48	Heart	Normal	H1	F	26	Thyroid	Normal
D2	F	34	Heart	Normal	H2	F	37	Thyroid	Normal
D3	F	52	Kidney, cortex	Normal	H3	F	51	Thyroid	Normal
D4	M	23	Kidney, cortex	Normal	H4	M	46	Tonsil	Normal
D5	M	63	Kidney, cortex	Normal	H5	M	13	Tonsil	Normal
D6	M	52	Kidney, medulla	Normal	H6	F	36	Uterus, cervix	Normal
D7	M	23	Kidney, medulla	Normal	H7	F	35	Uterus, cervix	Normal
D8	M	59	Peripheral nerve	Schwannoma	H8	F	44	Uterus, cervix	Normal
D9	M	29	Peripheral nerve	Schwannoma	H9	F	38	Uterus, endometrium	Normal
D10	F	44	Ureter	Normal	H10	F	41	Uterus, endometrium	Normal
D11	F	42	Ureter	Normal	H11	F	46	Uterus, endometrium	Normal
D12	F	57	Liver	Normal	H12	M	58	Skin	Malignant melanoma (tissue marker)
D13	F	57	Liver	Normal	H13	M	58	Skin	Malignant melanoma (tissue marker)

Fig. 133B

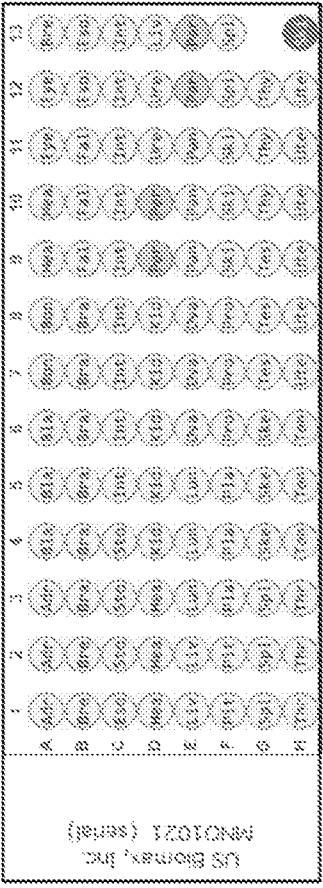


Figure 133A-133C



PSMGFR-antibody 18B4 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 134A  
Normal Adrenal Gland

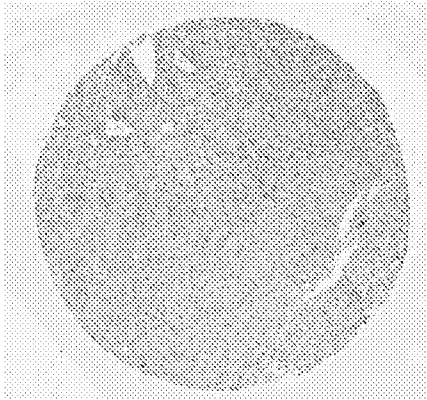


Fig. 134E

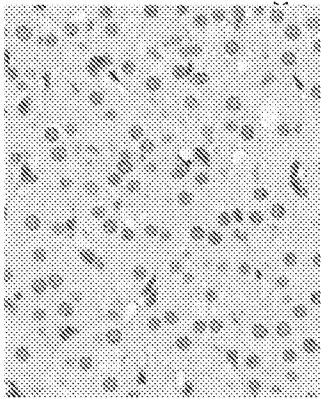


Fig. 134B  
Normal Breast

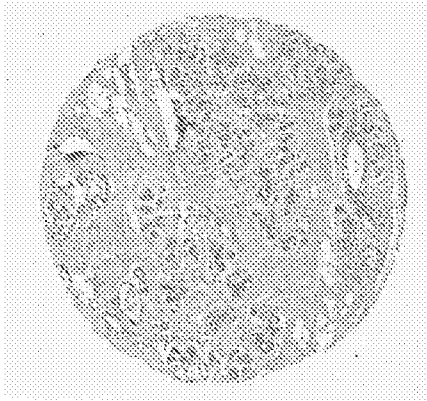


Fig. 134F



Fig. 134C  
Normal Fallopian Tubes

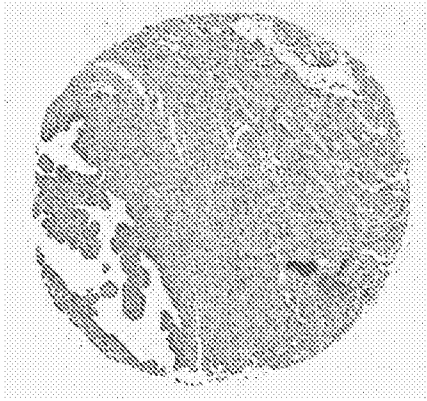


Fig. 134G

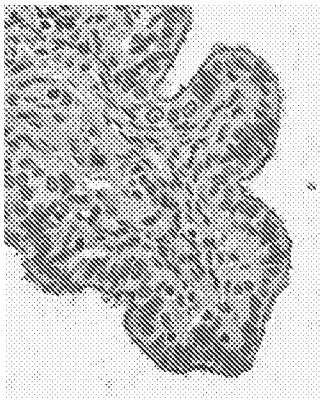


Fig. 134D  
Normal Kidney

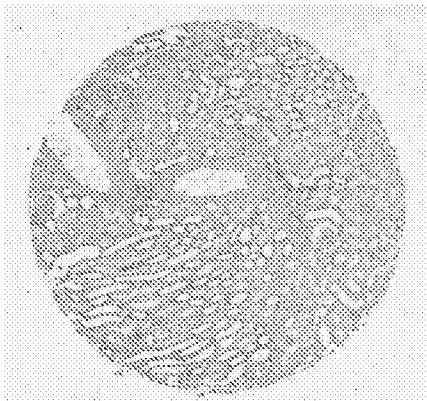


Fig. 134H

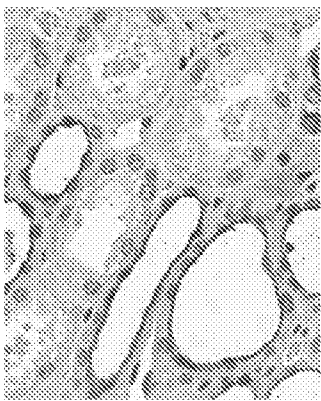


Figure 134A-134H

PSMGFR-antibody 18B4 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 134I  
Normal Heart

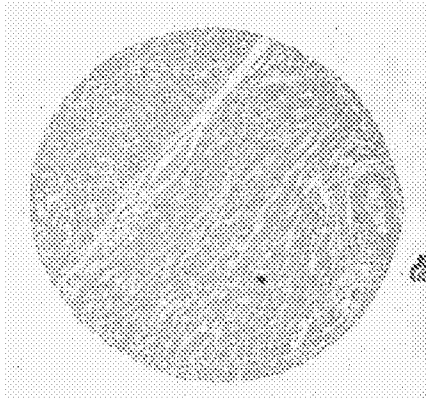


Fig. 134J  
Normal Liver

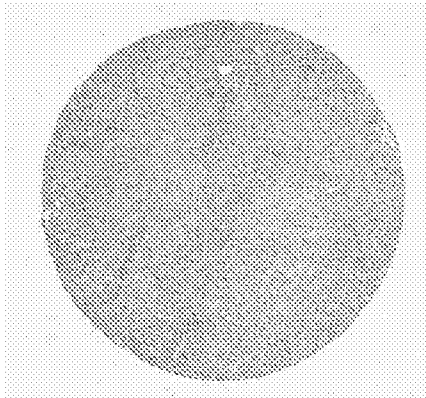


Fig. 134K  
Normal Lung

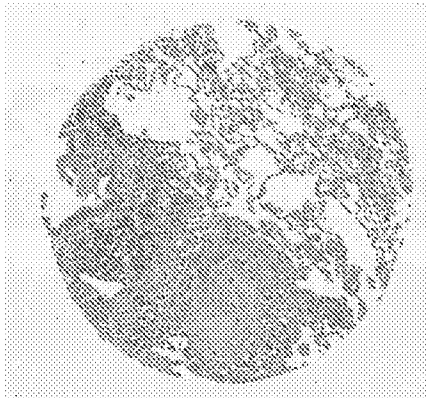


Fig. 134L  
Normal Ureter

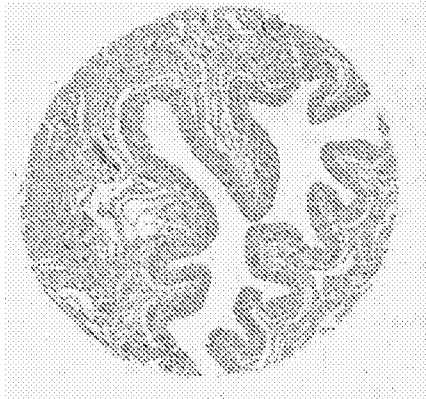


Fig. 134M

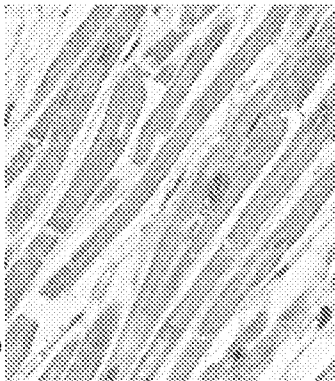


Fig. 134N

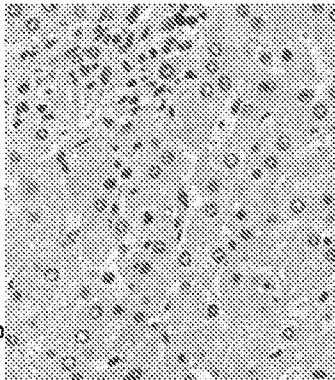


Fig. 134O

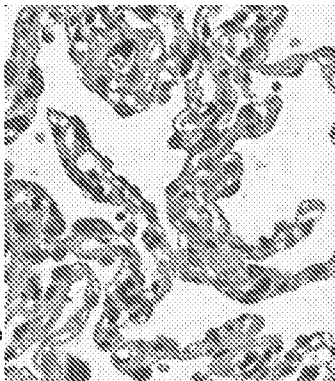


Fig. 134P

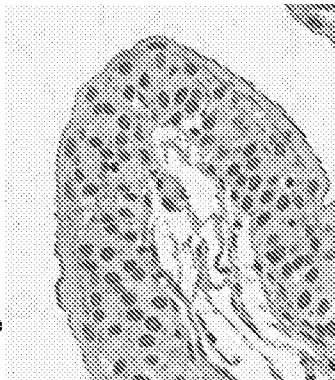


Figure 134I-134P

PSMGFR-antibody 18B4 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 134Q  
Normal eye

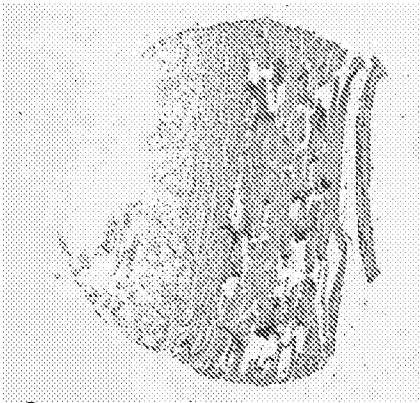


Fig. 134R  
Normal Cerebral cortex

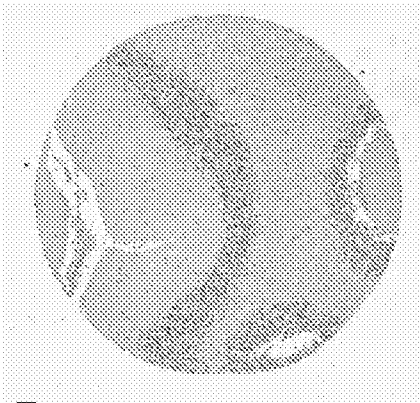


Fig. 134S  
Normal Bone marrow

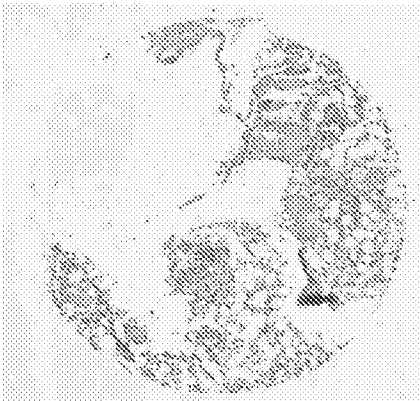


Fig. 134T  
Normal Skeletal muscle

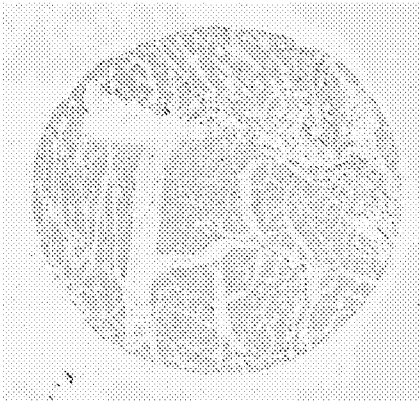


Fig. 134U



Fig. 134V

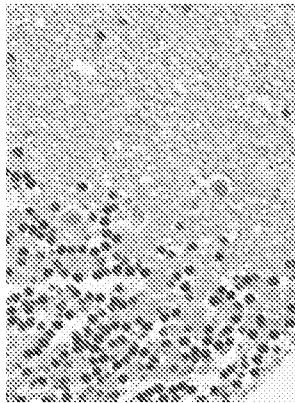


Fig. 134W

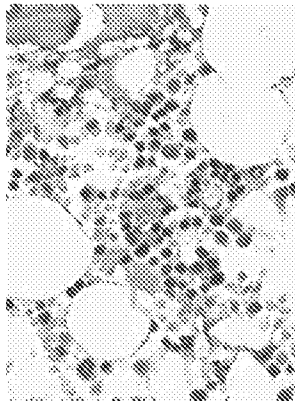


Fig. 134X



Figure 134Q-134X

The figure is a 10x10 grid of 100 circular images, each showing a different stage of chick embryo development. The grid is labeled with numbers 1 to 10 on both the left and top axes. The images show the progression from a single cell to a fully formed chick with a broken eggshell.

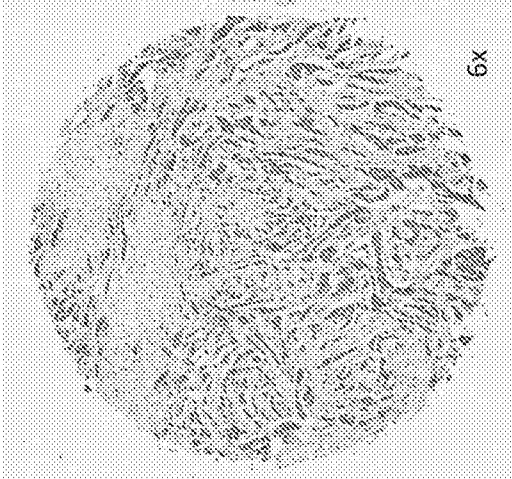
Figure 1 shows a 10x10 grid of 100 circular cells. The cells are arranged in 10 rows and 10 columns. The cells are labeled with numbers 1 through 100. The cells are arranged in a grid pattern. The cells are labeled with numbers 1 through 100.

Figure 135A-135C

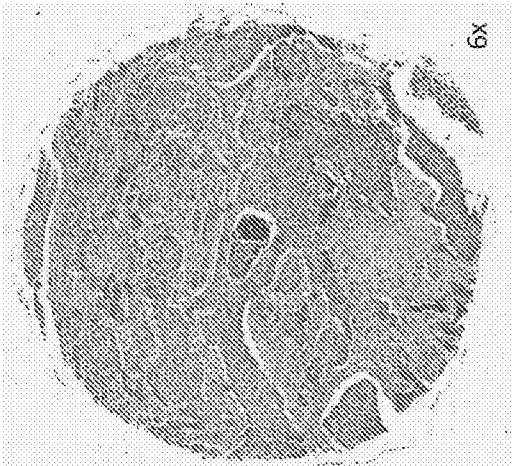
[illegible]

**PSMGFR-antibody 18B4 10 ug/mL**  
**Breast cancer tissue array BR1141**

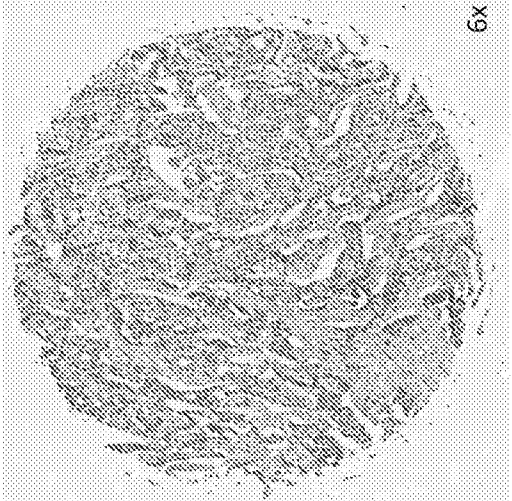
**Fig. 136A**  
Position: A4  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



**Fig. 136B**  
Position: G9  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T1N0M0



**Fig. 136C**  
Position: G11  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N1M0



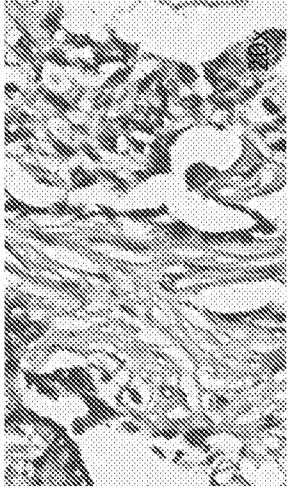
**Fig. 136D**



**Fig. 136E**



**Fig. 136F**



**Fig. 136A–136F**

PSMGFR antibody 18B4 IgG 10 ug/ml  
Esophageal cancer tissue array BC001113

Fig. 137A

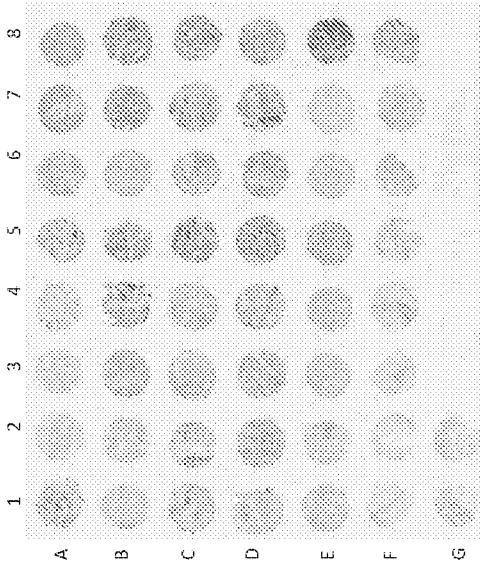


Fig. 137C

Position	Age	Sex	Organ/Anatomic	Pathology/diagnosis	Grade/Type	Position	Age	Sex	Organ/Anatomic	Pathology/diagnosis	Grade/Type
A1	47	F	Esophagus	Adenocarcinoma (solid)	II	D1	69	M	Cardia	Adenocarc	II
A2	33	F	Esophagus	Adenocarcinoma	II	D2	62	M	Cardia	Adenocarc	II
A3	68	F	Esophagus	Vuinous adenocarcinoma	I	D3	65	M	Cardia	Adenocarc	II
A4	80	M	Esophagus	Adenocarcinoma	II	D4	72	M	Cardia	Adenocarc	II
A5	61	M	Esophagus	Adenocarcinoma	II	D5	74	M	Stomach	Adenocarc	II
A6	57	M	Esophagus	Adenocarcinoma	II	D6	51	F	Stomach	Undifferent	V
A7	81	F	Esophagus	Undifferentiated adenocarc	IV	D7	51	F	Stomach	Undifferent	IV
A8	51	M	Esophagus	Adenocarcinoma	IV	D8	53	M	Stomach	Undifferent	IV
B1	69	M	Esophagus	Undifferentiated adenocarc	IV	E1	64	M	Cardia	Adenocarc	II
B2	97	F	Esophagus	Undifferentiated adenocarc	IV	E2	70	M	Cardia	Adenocarc	II
B3	72	M	Cardia	Adenocarcinoma	II	E3	73	M	Cardia	Undifferent	IV
B4	35	M	Cardia	Adenocarcinoma	I	E4	50	M	Cardia	Adenocarc	II
B5	55	F	Cardia	Adenocarcinoma	II	E5	84	M	Cardia	Adenocarc	II
B6	82	M	Cardia	Adenocarcinoma	II	E6	54	F	Cardia	Cancer ad	-
B7	64	F	Cardia	Undifferentiated adenocarc	IV	E7	43	M	Cardia	Adenocarc	I
B8	67	M	Cardia	Adenocarcinoma	I	E8	59	M	Cardia	Adenocarc	I
C1	80	F	Cardia	Adenocarcinoma	I	F1	64	F	Esophagus/Adjacent n	-	normal
C2	66	M	Cardia	Adenocarcinoma	II	F2	59	M	Esophagus/Adjacent n	-	normal
C3	63	F	Cardia	Adenocarcinoma	II	F3	67	M	Esophagus/Adjacent n	-	normal
C4	41	M	Cardia	Adenocarcinoma	II	F4	53	M	Esophagus/Adjacent n	-	normal
C5	70	M	Stomach	Adenocarcinoma	II	F5	43	M	Esophagus/Adjacent n	-	normal
C6	30	F	Stomach	Adenocarcinoma	I-II	F6	64	M	Cardia	Adjacent n	-
C7	61	M	Cardia	Adenocarcinoma	I-II	F7	71	M	Cardia	Adjacent n	-
C8	64	M	Cardia	Adenocarcinoma	II	F8	66	M	Esophagus/Adjacent n	-	normal
						G1	61	M	Cardia	Adjacent n	-
						G2	56	F	Cardia	Adjacent n	-

Fig. 137B

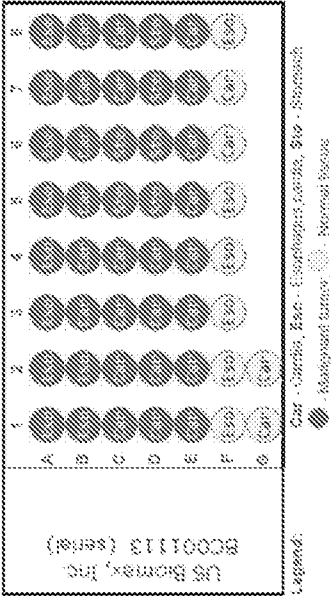
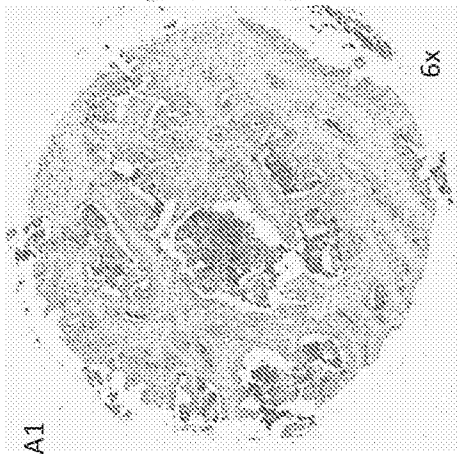


Figure 137A-137C



**PSMGFR antibody 18B4 IgG 10 ug/ml**  
**Esophageal cancer tissue array BC001113**

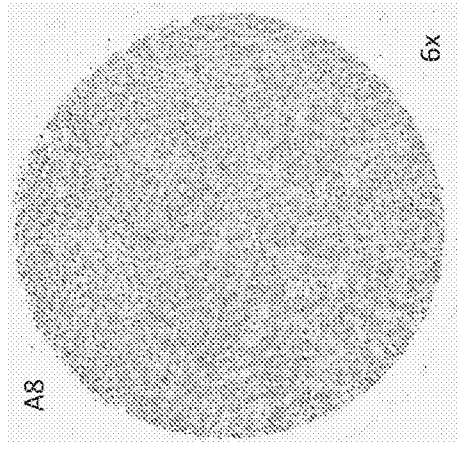
**Fig. 138A**  
**Esophageal cancer**



**Fig. 138B**  
**Esophageal cancer**



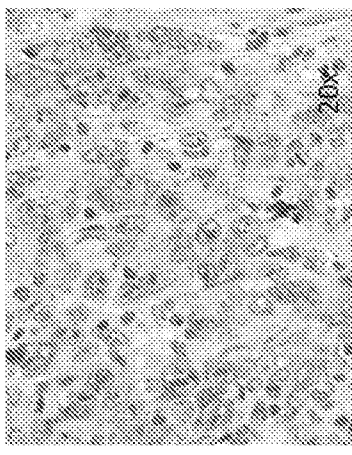
**Fig. 138C**  
**Esophageal cancer**



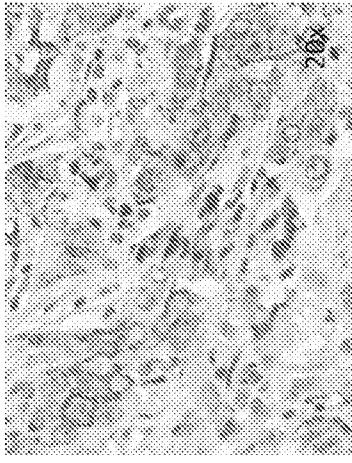
**Fig. 138D**



**Fig. 138E**



**Fig. 138F**



**Fig. 138A-138F**

PSMGFR-antibody 18G12 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 139A

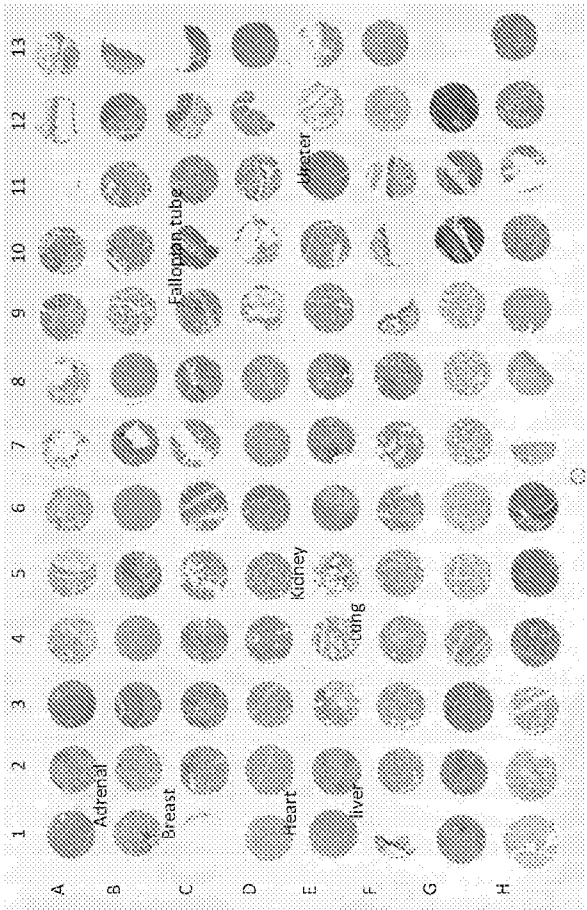


Fig. 139C

Position	Age	Sex	Organ/Anatomic Site	Pathology	Position	Age	Sex	Organ/Anatomic Site	Pathology
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	51	Adrenal gland	Normal	E3	F	72	Lung	Normal
A4	M	72	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	45	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	52	Bladder, urinary	Normal	E7	F	46	Ovary	Normal
A8	M	53	Bone, bone marrow	Normal	E8	F	46	Ovary	Normal
A9	F	44	Bone, bone marrow	Normal	E9	M	42	Pancreas	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	26	Pancreas	Normal
A11	M	18	Head and neck, salivary gland	Normal	E11	F	59	Pancreas	Normal
A12	M	50	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	F	35	Breast	Normal	E13	F	72	Parathyroid	Adenoma
B1	F	38	Breast	Normal	F1	F	3	Mammary gland	Normal
B2	F	38	Breast	Normal	F2	F	32	Pituitary gland	Normal
B3	M	58	Brain, cerebellum	Normal	F3	F	30	Pituitary gland	Normal
B4	F	58	Brain, cerebellum	Normal	F4	F	27	Pituitary gland	Normal
B5	M	58	Brain, cerebral cortex	Normal	F5	M	64	Prostate	Normal
B6	M	58	Brain, cerebral cortex	Normal	F6	M	65	Prostate	Normal
B7	F	42	Brain, cerebral cortex	Normal	F7	M	65	Prostate	Normal
B8	F	42	Brain, cerebral cortex	Normal	F8	M	32	Prostate	Normal
B9	F	32	Fallopian tube	Normal	F9	F	32	Skin	Normal
B10	F	32	Fallopian tube	Normal	F10	M	26	Skin	Normal
B11	F	24	Fallopian tube	Normal	F11	M	26	Skin	Normal
B12	M	34	Esophagus	Normal	F12	M	58	Spinal cord	Normal
B13	M	34	Esophagus	Normal	G1	M	27	Spleen	Normal
C1	M	45	Esophagus	Normal	G2	M	31	Spleen	Normal
C2	M	45	Esophagus	Normal	G3	M	31	Spleen	Normal
C3	M	77	Stomach	Normal	G4	F	60	Skeletal muscle	Normal
C4	M	77	Stomach	Normal	G5	M	49	Skeletal muscle	Normal
C5	M	45	Intestine, small intestine	Normal	G6	M	70	Skeletal muscle	Normal
C6	F	75	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C7	F	75	Intestine, small intestine	Normal	G8	M	43	Testis	Normal
C8	M	2	Intestine, colon	Normal	G9	M	43	Testis	Normal
C9	M	46	Intestine, colon	Normal	G10	M	15	Thymus	Normal
C10	M	74	Intestine, colon	Normal	G11	M	26	Thymus	Normal
C11	F	77	Intestine, rectum	Normal	G12	F	9	Thymus	Normal
C12	F	77	Intestine, rectum	Normal	G13	F	26	Thyroid	Normal
C13	M	26	Intestine, rectum	Normal	H1	F	37	Thyroid	Normal
D1	M	26	Heart	Normal	H2	F	37	Thyroid	Normal
D2	M	45	Heart	Normal	H3	F	51	Thyroid	Normal
D3	F	44	Heart	Normal	H4	M	46	Tonsil	Normal
D4	F	52	Kidney, cortex	Normal	H5	M	37	Tonsil	Normal
D5	M	23	Kidney, cortex	Normal	H6	M	13	Tonsil	Normal
D6	M	53	Kidney, medulla	Normal	H7	F	36	Uterus, cervix	Normal
D7	M	53	Kidney, medulla	Normal	H8	F	35	Uterus, cervix	Normal
D8	M	53	Kidney, medulla	Schwannoma	H9	F	44	Uterus, cervix	Normal
D9	M	39	Peripheral nerve	Schwannoma	H10	F	36	Uterus	Normal
D10	F	28	Peripheral nerve	Schwannoma	H11	F	41	Uterus	Normal
D11	F	44	Ureter	Normal	H12	F	46	Uterus	Normal
D12	F	42	Ureter	Normal	H13	M	56	Skin	Malignant melanoma (tissue marker)
D13	F	57	Liver	Normal					

Fig. 139B

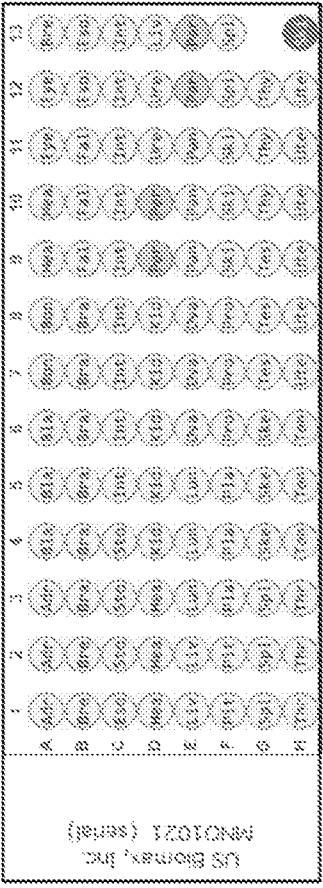


Figure 139A-139C



PSMGFR-antibody 18G12 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 140A  
Normal Adrenal Gland

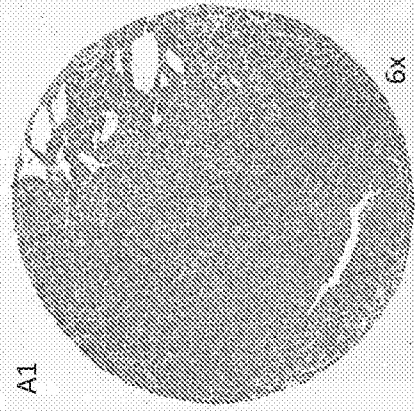


Fig. 140B  
Normal Breast

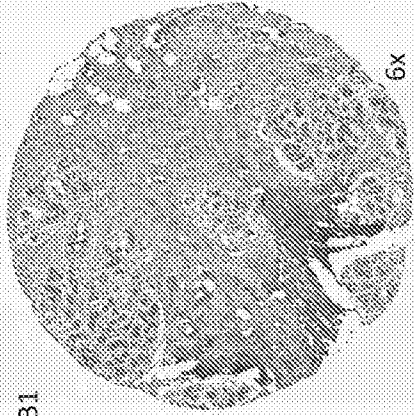


Fig. 140C  
Normal Fallopian Tubes

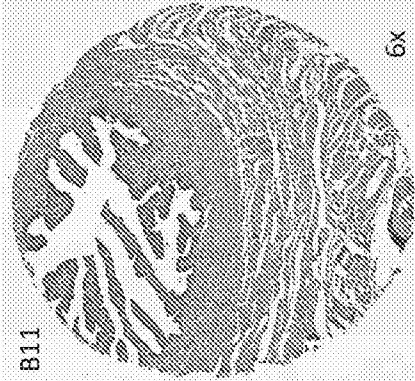


Fig. 140D  
Normal Kidney

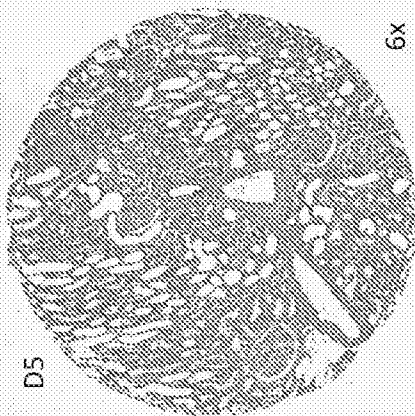


Fig. 140E

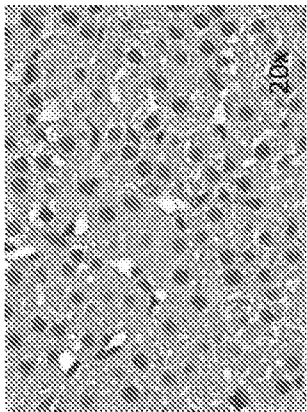


Fig. 140F

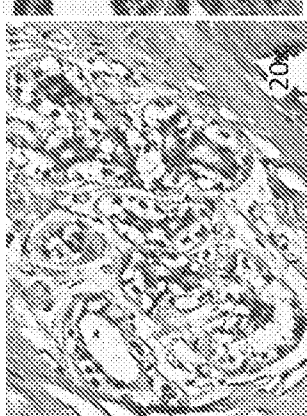


Fig. 140G

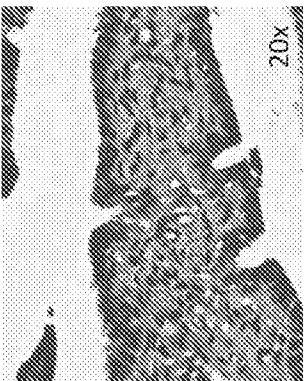


Fig. 140H

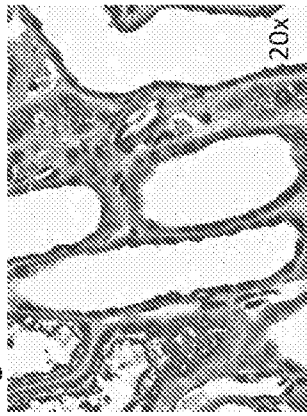
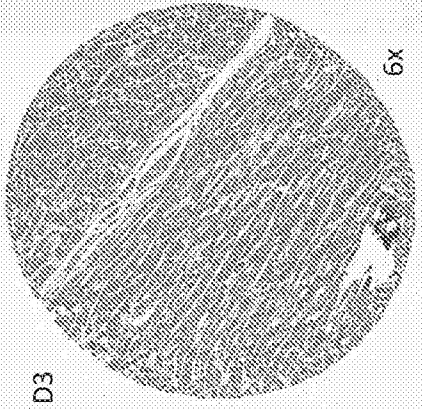


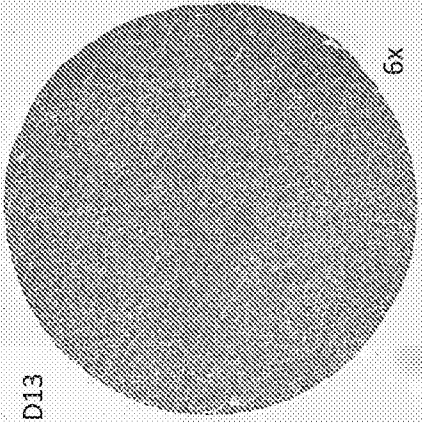
Figure 140A-140H

**PSMGFR-antibody 18G12 IgG 10 ug/mL**  
**FDA Normal tissue array MNO1021**

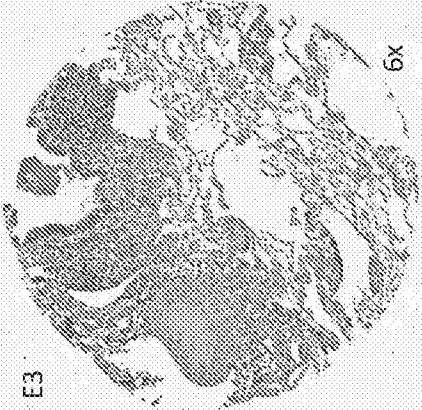
**Fig. 140I**  
**Normal Heart**



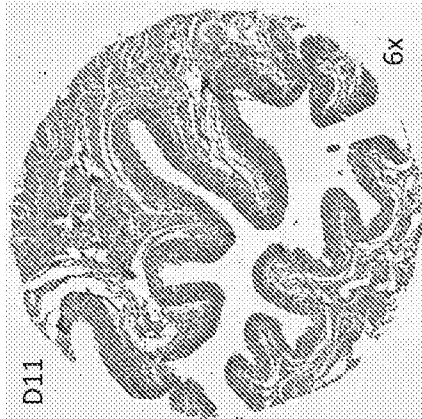
**Fig. 140J**  
**Normal Liver**



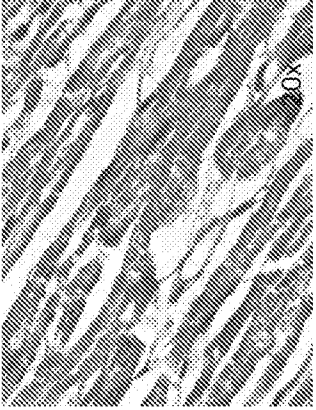
**Fig. 140K**  
**Normal Lung**



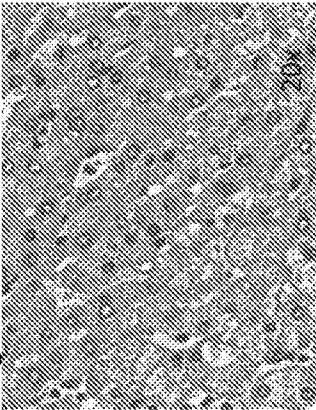
**Fig. 140L**  
**Normal Ureter**



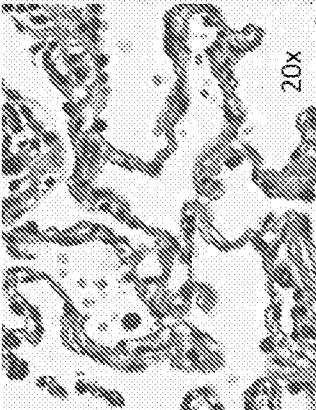
**Fig. 140M**



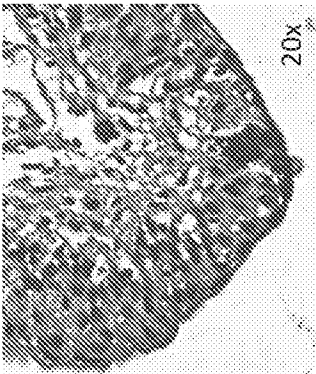
**Fig. 140N**



**Fig. 140O**



**Fig. 140P**



**Figure 140I-140P**

PSMGFR-antibody 18G12 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 140Q  
Normal eye

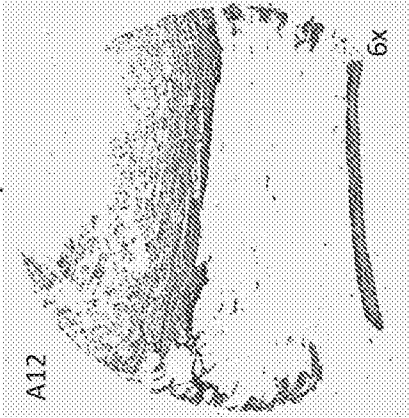


Fig. 140R  
Normal Cerebral cortex

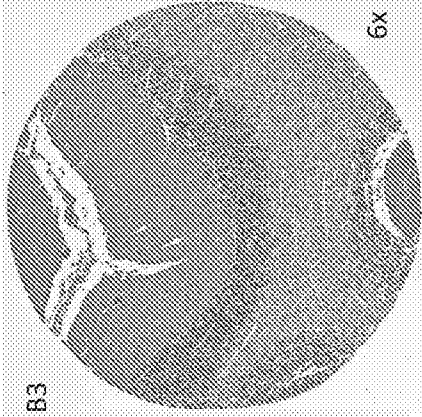


Fig. 140S  
Normal Bone marrow

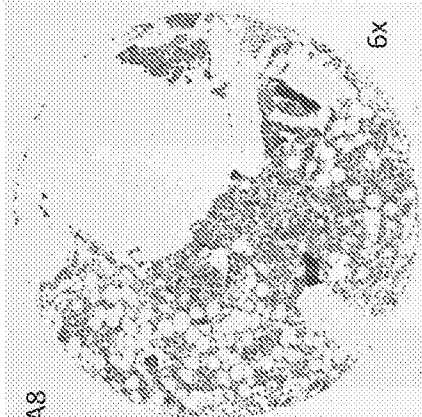


Fig. 140T  
Normal Skeletal muscle

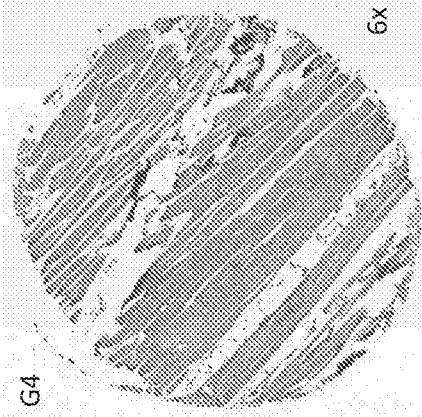


Fig. 140U

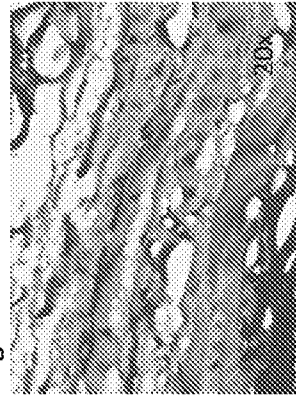


Fig. 140V

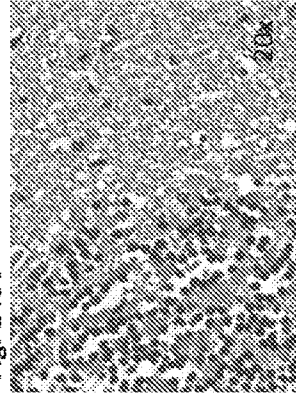


Fig. 140W



Fig. 140X

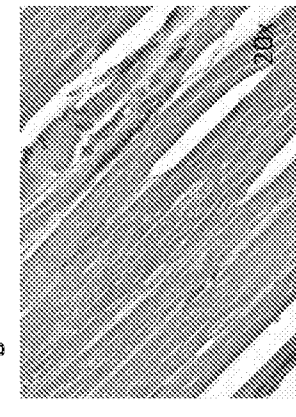


Figure 140Q-140X



PSMGFR-antibody 18G12 15 ug/mL  
Breast cancer tissue array BR1141

Fig. 142A

Position: C5  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0

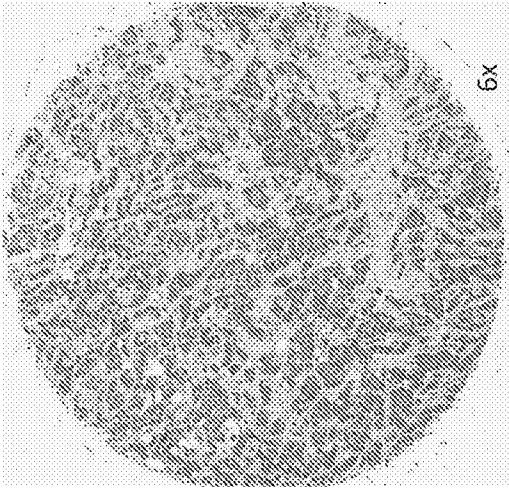


Fig. 142B

Position: G9  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T1N0M0

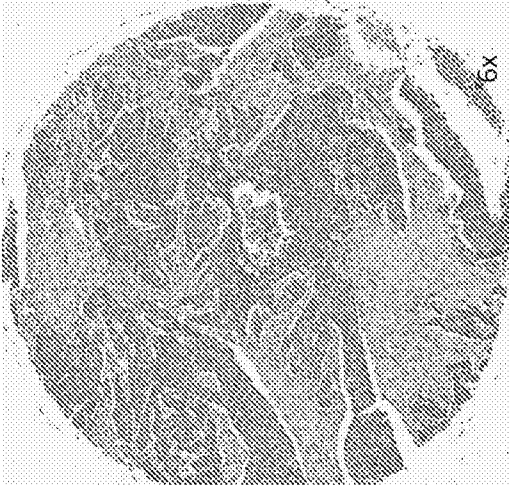


Fig. 142C

Position: I1  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N1M0

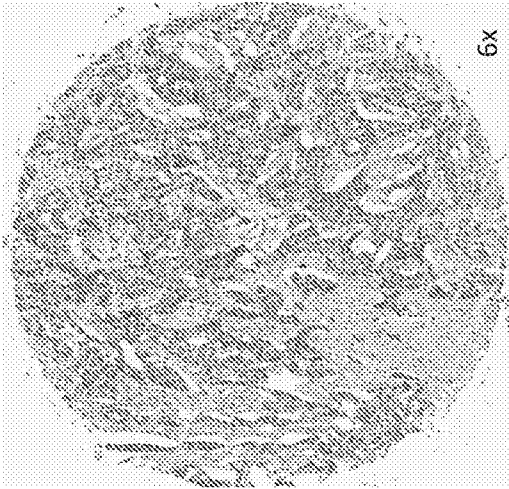


Fig. 142D

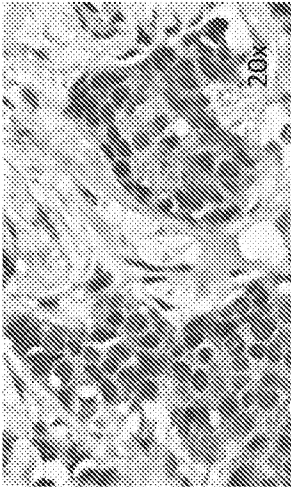


Fig. 142E

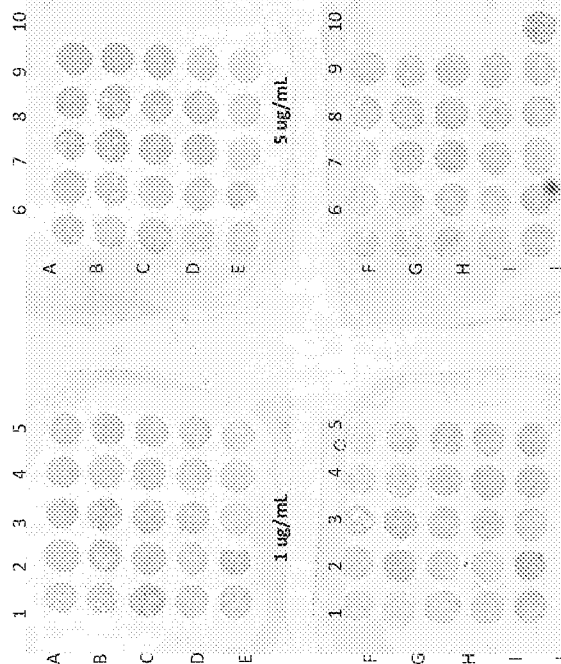


Fig. 142F



Fig. 142A-142F





3345

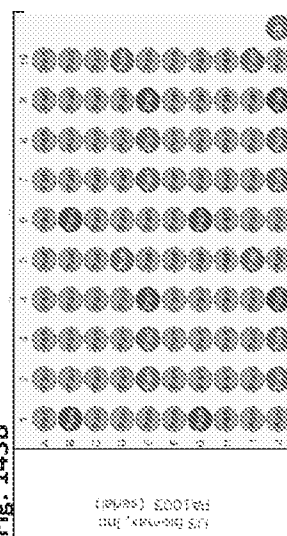


Fig. 143A - 143C

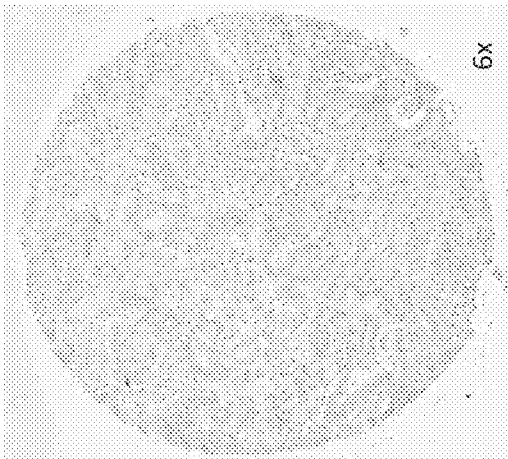
File 143C

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**PSMGFR-antibody 18G12 15 ug/ml**  
**Pancreatic cancer tissue array PA1003**

**Fig. 144A**

Position: A2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0

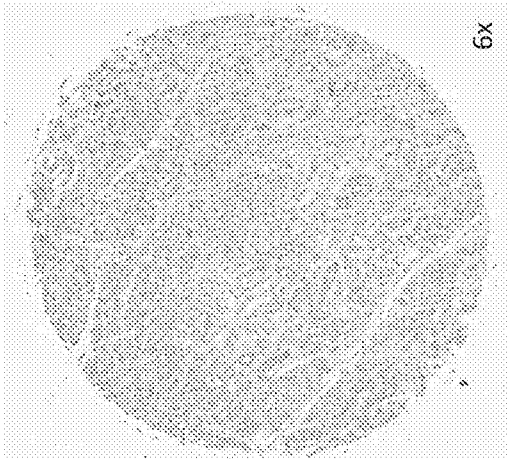


**Fig. 144D**

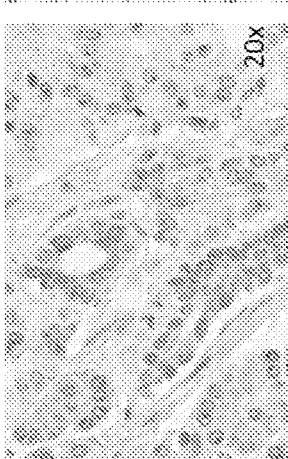


**Fig. 144B**

Position: B2  
Cell Type: Adenocarcinoma  
Tumor Grade: 1  
TNM: T3N0M0

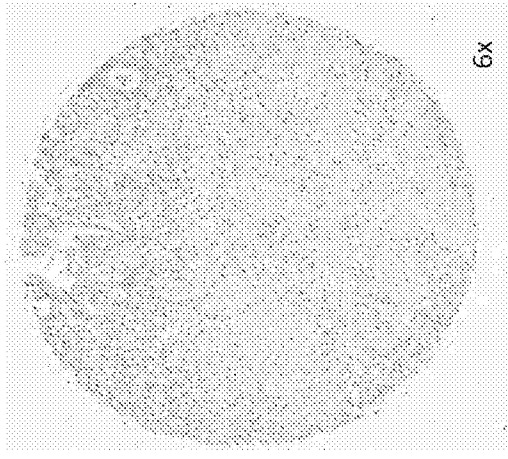


**Fig. 144E**

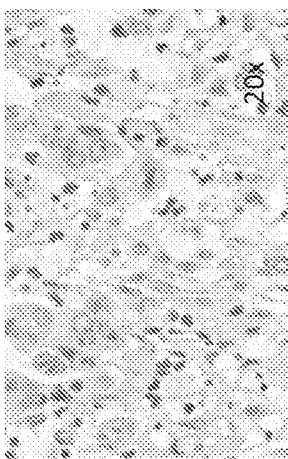


**Fig. 144C**

Position: C2  
Cell Type: Adenocarcinoma  
Tumor Grade: 1  
TNM: T3N0M0



**Fig. 144F**



**Fig. 144A-144F**

PSMGFR antibody 18G12 IgG 30 ug/ml  
Esophageal cancer tissue array BC001113

Fig. 145A

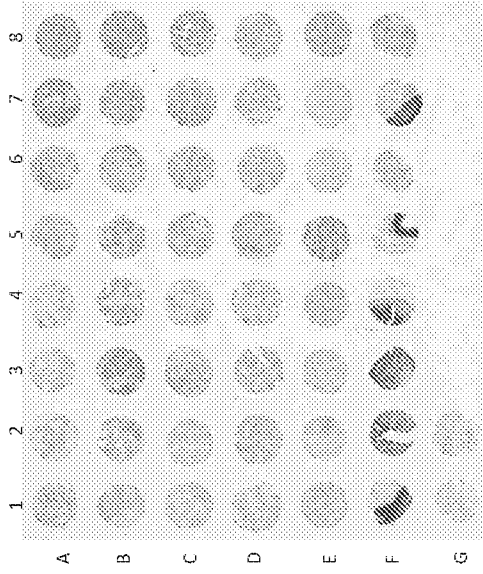


Fig. 145C

Position	Age	Sex	Organ/Anatomic	Pathology	Diagnosis	Grade	Type	Position	Age	Sex	Organ/Anatomic	Pathology	Diagnosis	Grade	Type
A1	47	F	Esophagus	Adenocarcinoma	(sho)	II	malignant	D1	69	M	Cardia	Adenocarc			malignant
A2	33	F	Esophagus	Adenocarcinoma		II	malignant	D2	62	M	Cardia	Adenocarc			malignant
A3	60	F	Esophagus	Vuinous adenocarcinoma		I	malignant	D3	65	M	Cardia	Adenocarc			malignant
A4	80	M	Esophagus	Adenocarcinoma		II	malignant	D4	72	M	Cardia	Adenocarc			malignant
A5	61	M	Esophagus	Adenocarcinoma		II	malignant	D5	74	M	Stomach	Adenocarc			malignant
A6	57	M	Esophagus	Adenocarcinoma		II	malignant	D6	51	F	Stomach	Undifferent			malignant
A7	81	M	Esophagus	Undifferentiated adenocarc		IV	malignant	D7	51	F	Stomach	Undifferent			malignant
A8	51	F	Esophagus	Adenocarcinoma		IV	malignant	D8	53	M	Stomach	Undifferent			malignant
B1	69	M	Esophagus	Undifferentiated adenocarc		IV	malignant	E1	64	M	Cardia	Adenocarc			malignant
B2	97	F	Esophagus	Adenocarcinoma		IV	malignant	E2	70	M	Cardia	Adenocarc			malignant
B3	72	M	Cardia	Adenocarcinoma		I	malignant	E3	73	M	Cardia	Undifferent			malignant
B4	35	M	Cardia	Adenocarcinoma		I	malignant	E4	50	M	Cardia	Adenocarc			malignant
B5	55	F	Cardia	Adenocarcinoma		II	malignant	E5	84	M	Cardia	Adenocarc			malignant
B6	92	M	Cardia	Adenocarcinoma		II	malignant	E6	54	F	Cardia	Cancer ad			malignant
B7	64	F	Cardia	Undifferentiated adenocarc		IV	malignant	E7	43	M	Cardia	Adenocarc			malignant
B8	87	M	Cardia	Adenocarcinoma		I	malignant	E8	59	M	Cardia	Adenocarc			malignant
C1	80	F	Cardia	Adenocarcinoma		I	malignant	F1	64	F	Esophagus/Adjacent n			normal	
C2	66	M	Cardia	Adenocarcinoma		II	malignant	F2	59	M	Esophagus/Adjacent n			normal	
C3	63	F	Cardia	Adenocarcinoma		II	malignant	F3	67	M	Esophagus/Adjacent n			normal	
C4	41	M	Cardia	Adenocarcinoma		II	malignant	F4	53	M	Esophagus/Adjacent n			normal	
C5	70	M	Stomach	Adenocarcinoma		II	malignant	F5	43	M	Esophagus/Adjacent n			normal	
C6	30	F	Stomach	Adenocarcinoma		I-II	malignant	F6	64	M	Cardia	Adjacent n			normal
C7	61	M	Cardia	Adenocarcinoma		I-II	malignant	F7	71	M	Cardia	Adjacent n			normal
C8	64	M	Cardia	Adenocarcinoma		II	malignant	F8	66	M	Esophagus/Adjacent n			normal	
								G1	61	M	Cardia	Adjacent n			normal
								G2	56	F	Cardia	Adjacent n			normal

Fig. 145B

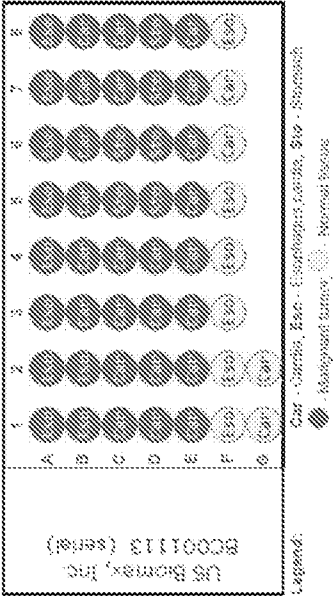
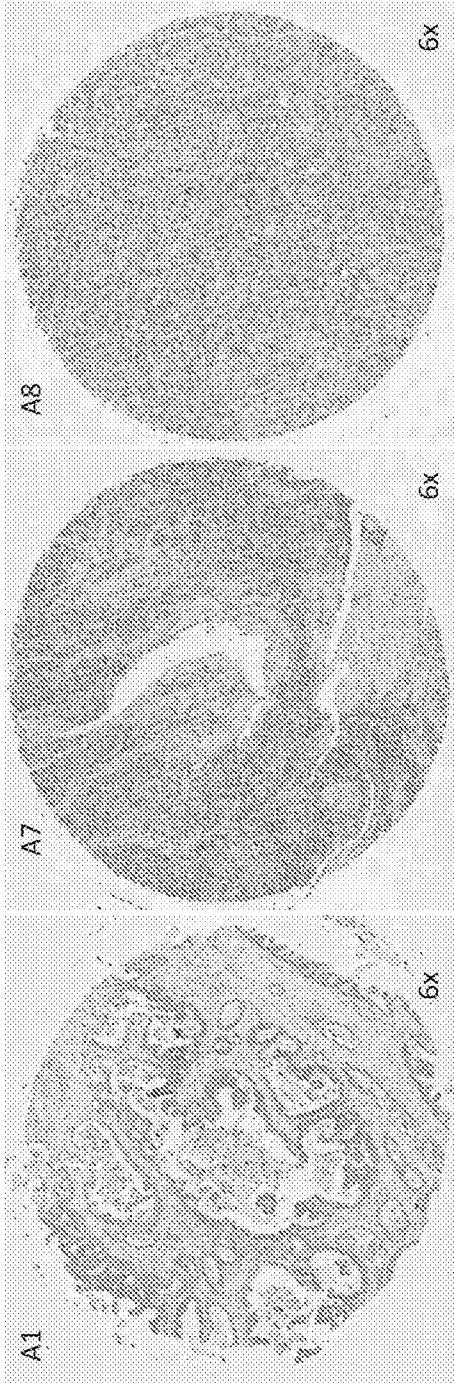


Figure 145A-145C



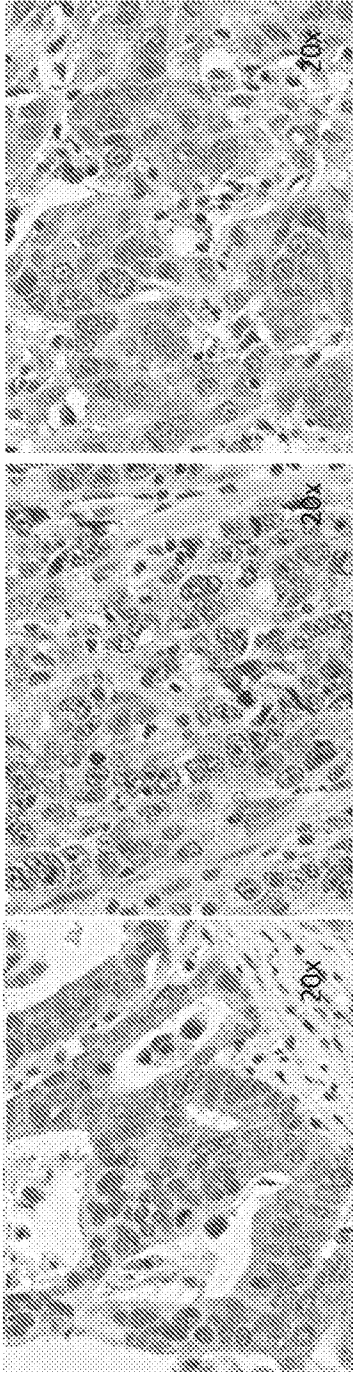
**PSMGFR antibody 18G12 IgG 30 ug/ml**  
**Esophageal cancer tissue array BC001113**



**Fig. 146A**  
**Esophageal cancer**

**Fig. 146B**  
**Esophageal cancer**

**Fig. 146C**  
**Esophageal cancer**



**Fig. 146D**

**Fig. 146E**

**Fig. 146F**

**Fig. 146A-146F**

PSMGFR-antibody 25E6 IgG 5.0 ug/mL  
FDA Normal tissue array MNO1021

Fig. 147A

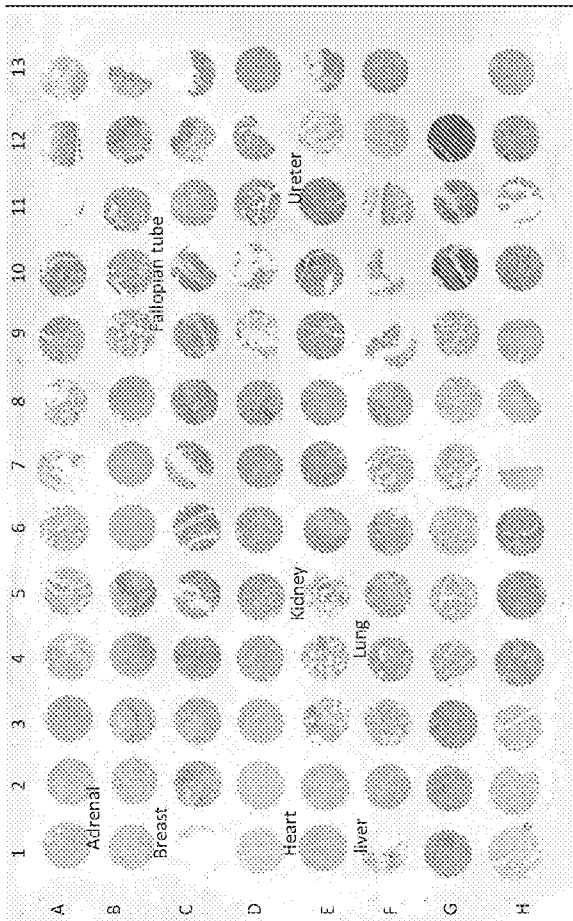


Fig. 147C

Position	Age	Sex	Organ/Anatomic Site	Pathology	Position	Age	Sex	Organ/Anatomic Site	Pathology
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	33	Adrenal gland	Normal	E3	M	72	Lung	Normal
A4	M	75	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	43	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bladder, urinary	Normal	E7	F	46	Ovary	Normal
A8	F	43	Bone, bone marrow	Normal	E8	F	40	Ovary	Normal
A9	M	44	Head and neck, salivary gland	Normal	E9	M	42	Pancreas	Normal
A10	M	76	Head and neck, salivary gland	Normal	E10	F	26	Pancreas	Normal
A11	M	35	Eye	Normal	E11	F	39	Pituitary gland	Normal
A12	M	30	Eye	Normal	E12	M	77	Pituitary gland	Adenoma
A13	F	38	Breast	Normal	E13	M	72	Pituitary gland	Adenoma
B1	F	38	Breast	Normal	F1	F	3	Pituitary gland	Normal
B2	F	33	Breast	Normal	F2	F	32	Pituitary gland	Normal
B3	M	56	Brain, cerebellum	Normal	F3	F	30	Pituitary gland	Normal
B4	M	65	Brain, cerebellum	Normal	F4	F	30	Pituitary gland	Normal
B5	F	58	Brain, cerebral cortex	Normal	F5	F	27	Pituitary gland	Normal
B6	M	58	Brain, cerebral cortex	Normal	F6	M	94	Prostate	Normal
B7	M	65	Brain, cerebral cortex	Normal	F7	M	95	Prostate	Normal
B8	F	42	Fallopian tube	Normal	F8	M	65	Prostate	Normal
B9	F	42	Fallopian tube	Normal	F9	F	32	Skin	Normal
B10	F	32	Fallopian tube	Normal	F10	F	86	Skin	Normal
B11	F	24	Fallopian tube	Normal	F11	M	26	Skin	Normal
B12	F	45	Esophagus	Normal	F12	M	47	Spinal cord	Normal
B13	F	66	Esophagus	Normal	F13	M	36	Spinal cord	Normal
C1	M	54	Esophagus	Normal	G1	M	27	Spleen	Normal
C2	M	45	Stomach	Normal	G2	M	90	Spleen	Normal
C3	M	49	Stomach	Normal	G3	M	31	Spleen	Normal
C4	M	77	Stomach	Normal	G4	F	90	Skeletal muscle	Normal
C5	M	45	Intestine, small intestine	Normal	G5	F	49	Skeletal muscle	Normal
C6	F	75	Intestine, small intestine	Normal	G6	M	70	Skeletal muscle	Normal
C7	F	75	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C8	M	2	Intestine, colon	Normal	G8	M	30	Testis	Normal
C9	M	46	Intestine, colon	Normal	G9	M	77	Testis	Normal
C10	M	74	Intestine, colon	Normal	G10	M	15	Thymus	Normal
C11	F	47	Intestine, rectum	Normal	G11	M	26	Thymus	Normal
C12	F	75	Intestine, rectum	Normal	G12	F	9	Thymus	Normal
C13	M	96	Intestine, rectum	Normal	G13	-	-	-	-
D1	M	56	Heart	Normal	H1	F	26	Thyroid	Normal
D2	M	46	Heart	Normal	H2	F	37	Thyroid	Normal
D3	F	34	Heart	Normal	H3	F	51	Thyroid	Normal
D4	F	52	Kidney, cortex	Normal	H4	M	46	Tonsil	Normal
D5	M	23	Kidney, cortex	Normal	H5	M	37	Tonsil	Normal
D6	M	63	Kidney, cortex	Normal	H6	M	13	Tonsil	Normal
D7	F	52	Kidney, medulla	Normal	H7	F	36	Uterus, cervix	Normal
D8	M	23	Kidney, medulla	Normal	H8	F	35	Uterus, cervix	Normal
D9	M	59	Peripheral nerve	Schwannoma	H9	F	44	Uterus, cervix	Normal
D10	F	29	Peripheral nerve	Schwannoma	H10	F	38	Uterus, endometrium	Normal
D11	F	44	Ureter	Normal	H11	F	41	Uterus, endometrium	Normal
D12	F	42	Ureter	Normal	H12	F	48	Uterus, endometrium	Normal
D13	F	57	Liver	Normal	H13	M	58	Skin	Malignant melanoma (tissue marker)

Fig. 147B

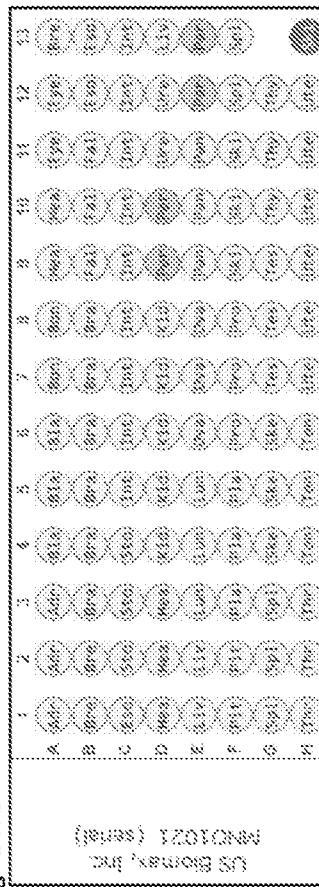


Figure 147A-147C

PSMGFR-antibody 25E6 IgG 5.0 ug/mL  
FDA Normal tissue array MNO1021

Fig. 148A

Normal Adrenal Gland

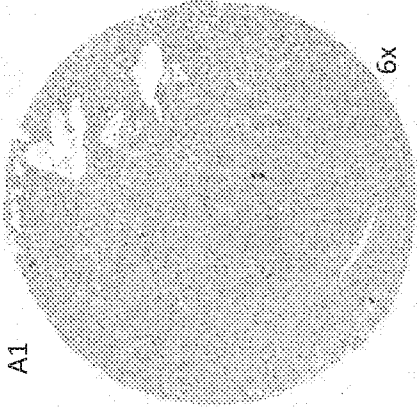


Fig. 148B

Normal Breast

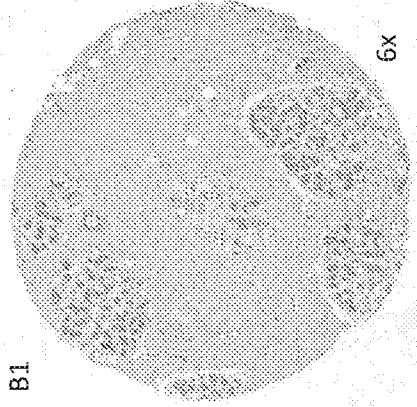


Fig. 148C

Normal Fallopian Tubes

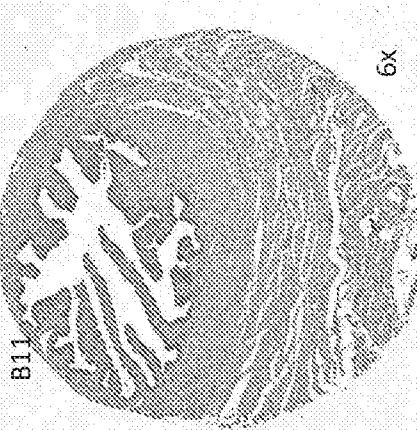


Fig. 148D

Normal Kidney

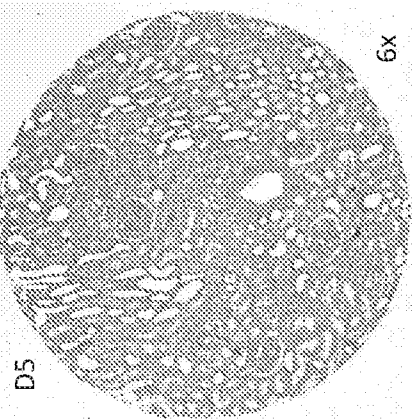


Fig. 148E

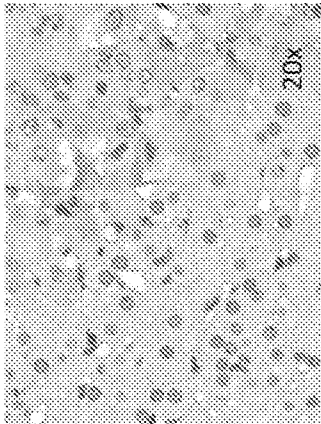


Fig. 148F



Fig. 148G

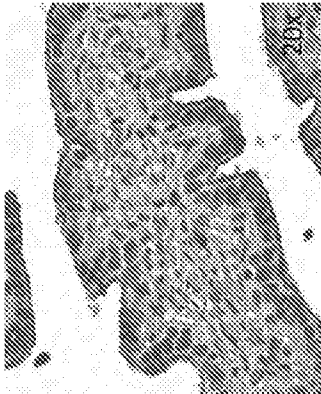


Fig. 148H



Figure 148A-148H

PSMGFR-antibody 25E6 IgG 5.0 ug/mL  
FDA Normal tissue array MNO1021

Fig. 148I  
Normal Heart

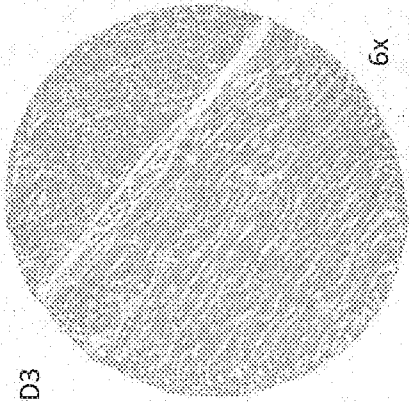


Fig. 148J  
Normal Liver

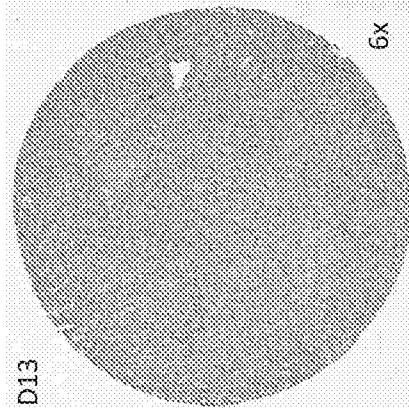


Fig. 148K  
Normal Lung

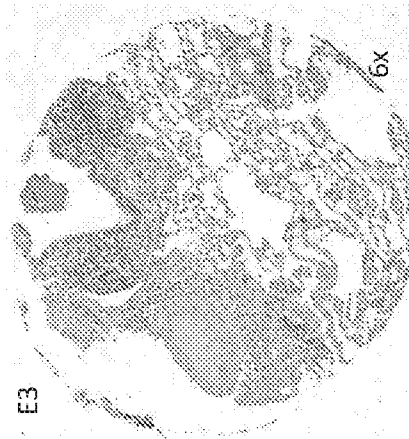


Fig. 148L  
Normal Ureter

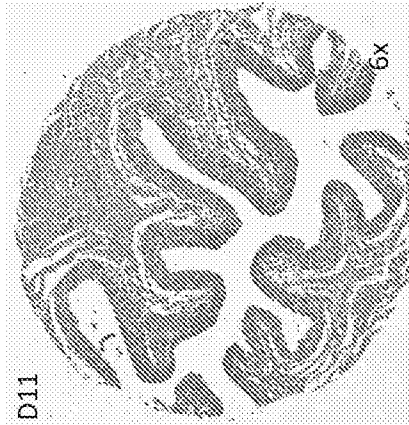


Fig. 148M

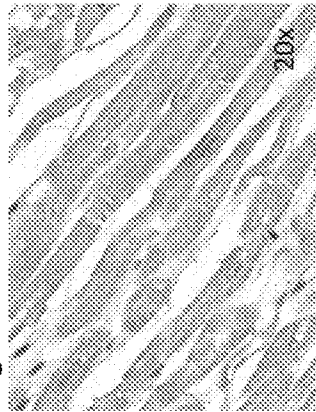


Fig. 148N

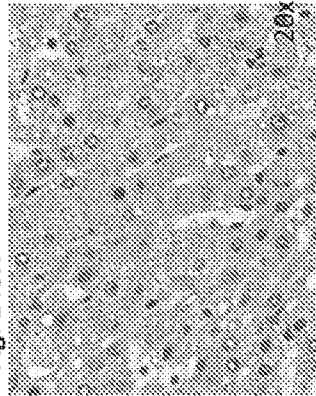


Fig. 148O



Fig. 148P

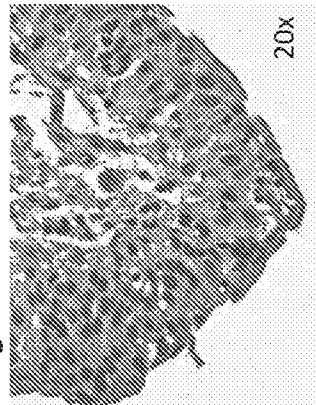


Figure 148I-148P

PSMGFR-antibody 25E6 IgG 5.0 ug/mL  
FDA Normal tissue array MNO1021

Fig. 148Q  
Normal eye

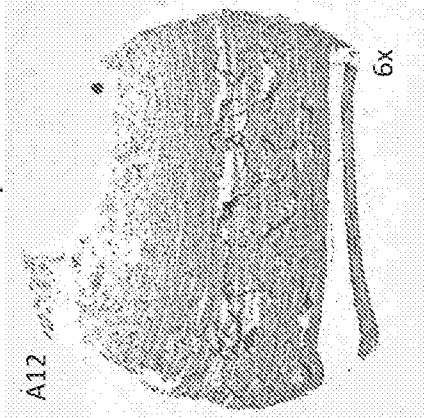


Fig. 148R  
Normal Cerebral cortex

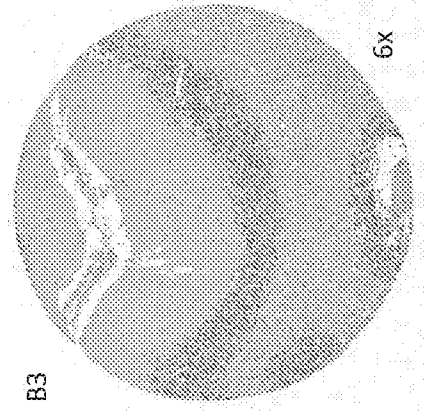


Fig. 148S  
Normal Bone marrow

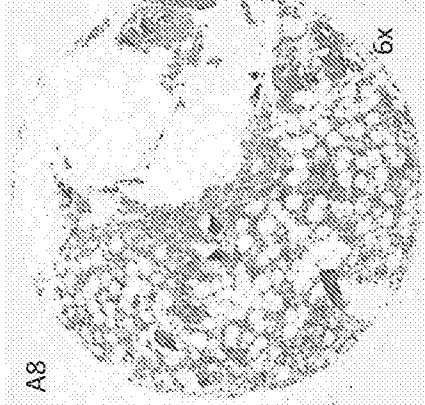


Fig. 148T  
Normal Skeletal muscle



Fig. 148U



Fig. 148V

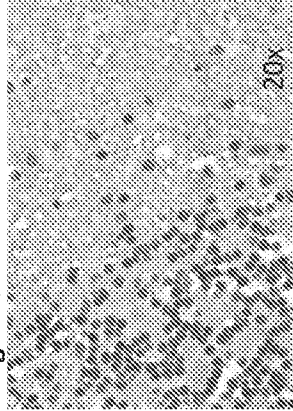


Fig. 148W



Fig. 148X

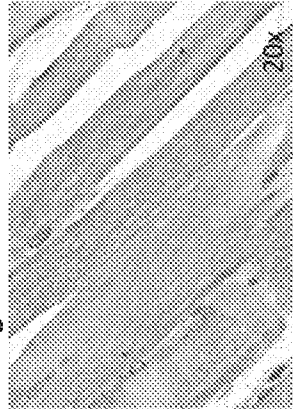


Figure 148Q-148X

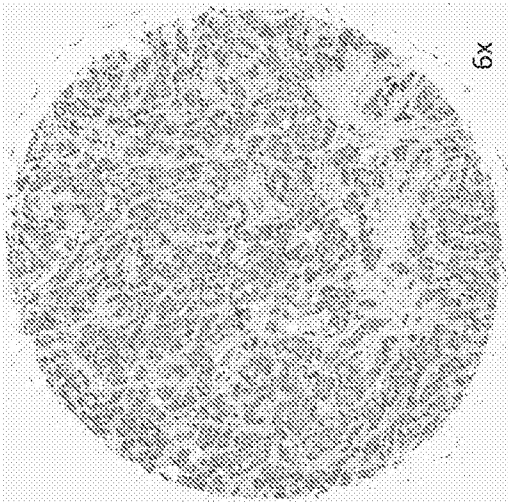




PSMGFR-antibody 25E6 5 ug/mL  
Breast cancer tissue array BR1141

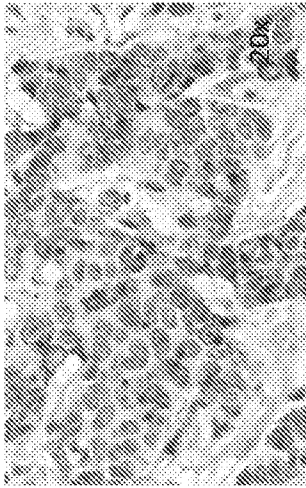
Fig. 150A

Position: C5  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



6x

Fig. 150D



20x

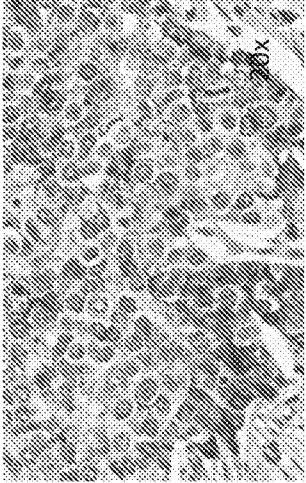
Fig. 150B

Position: G9  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T1N0M0



6x

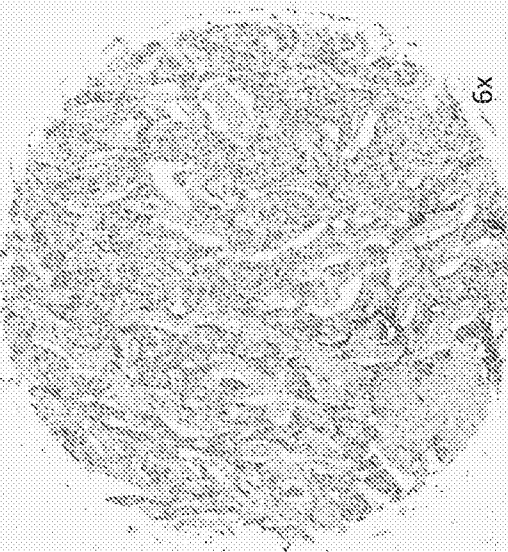
Fig. 150E



20x

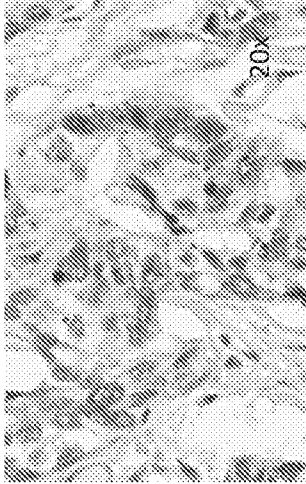
Fig. 150C

Position: I1  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N1M0



6x

Fig. 150F



20x

Fig. 150A-150F

PSMGFR-antibody 25E6 0.1-5 ug/ml  
Pancreatic cancer tissue array PA1003

Fig. 151A

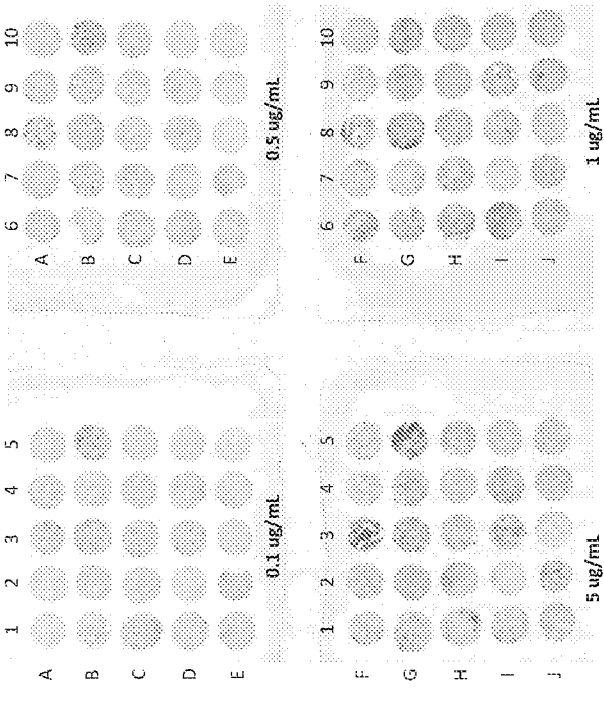
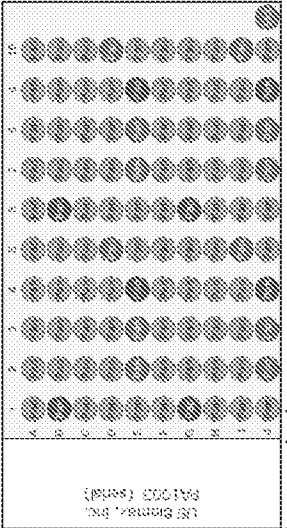


Fig. 151B



Legend:   
 0.1 ug/ml   
 0.5 ug/ml   
 1 ug/ml   
 25E6   
 PA1003 (panc)

Fig. 151A-151C

Fig. 151C

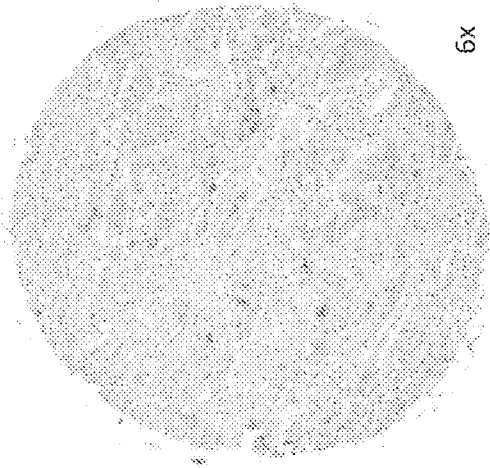
A1	38	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A2	41	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A3	50	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A4	65	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A5	53	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A6	39	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A7	44	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A8	59	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A9	65	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A10	59	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B1	42	M	Adenocarcinoma	TSN001	2	IV	malignant	TSN001	2	IV	malignant
B2	52	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B3	57	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B4	34	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B5	71	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B6	42	M	Adenocarcinoma	TSN001	2	IV	malignant	TSN001	2	IV	malignant
B7	52	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B8	57	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B9	34	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B10	72	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C1	47	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C2	64	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C3	51	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C4	68	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C5	41	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C6	47	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C7	64	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C8	51	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C9	68	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C10	41	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D1	58	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D2	41	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D3	77	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D4	50	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D5	56	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D6	58	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D7	41	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D8	72	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D9	50	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D10	50	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E1	53	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E2	60	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E3	50	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E4	40	F	Adenocarcinoma	TSN001	2	IV	malignant	TSN001	2	IV	malignant
E5	55	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E6	55	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E7	60	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E8	56	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E9	60	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E10	66	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant



**PSMGFR-antibody 25E6 0.1 ug/ml**  
**Pancreatic cancer tissue array PA1003**

**Fig. 152A**

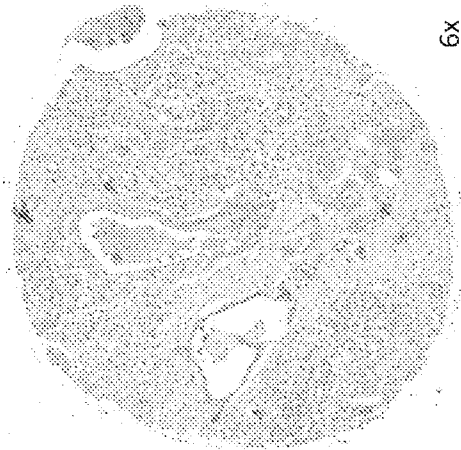
Position: A2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



6x

**Fig. 152B**

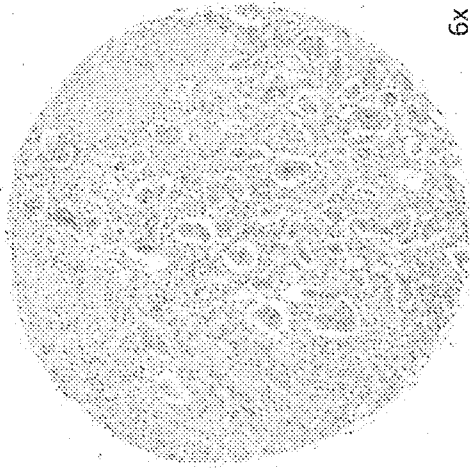
Position: B2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



6x

**Fig. 152C**

Position: C2  
Cell Type: Adenocarcinoma  
Tumor Grade: 1  
TNM: T3N0M0



6x

**Fig. 152D**



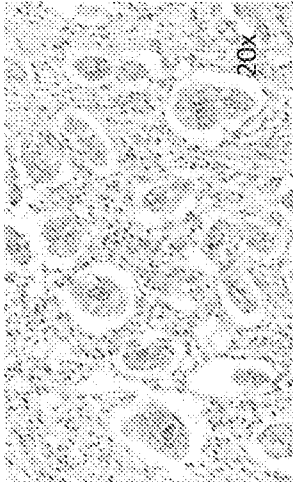
20x

**Fig. 152E**



20x

**Fig. 152F**



20x

**Fig. 152A-152F**

PSMGFR-antibody 28F9 IgG 15 ug/ml  
FDA Normal tissue array MNO1021

Fig. 153A

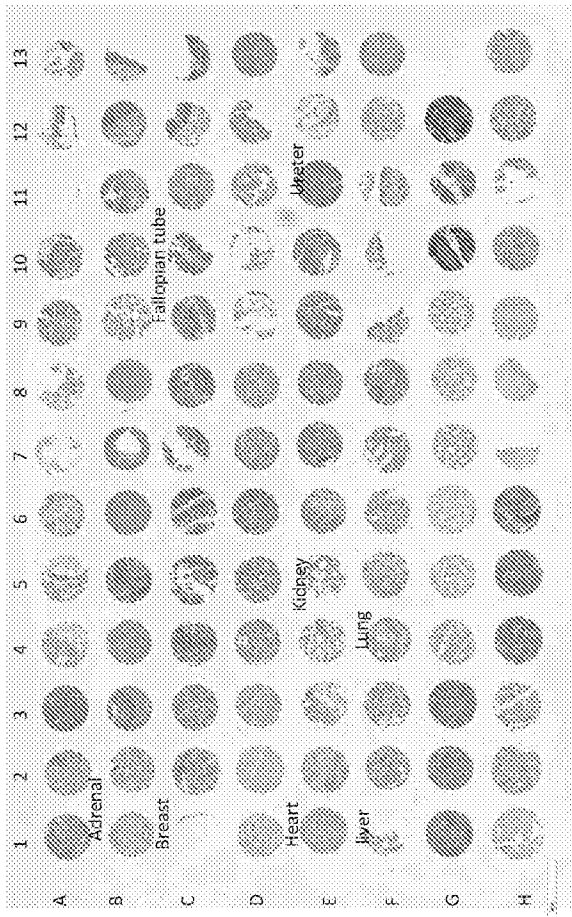


Fig. 153C

Position	Age	Sex	Organ/Anatomic Site	Pathology	Diagnosis	Position	Age	Sex	Organ/Anatomic Site	Pathology	Diagnosis	
A1	F	31	Adrenal gland	Normal	Normal	E1	M	59	Liver	Normal	Normal	
A2	M	51	Adrenal gland		Normal	E2	F	27	Liver	Normal	Normal	
A3	M	33	Adrenal gland		Normal	E3	M	72	Lung	Normal	Normal	
A4	M	75	Bladder, urinary		Normal	E4	F	55	Lung	Normal	Normal	
A5	M	72	Bladder, urinary		Normal	E5	M	43	Lung	Normal	Normal	
A6	M	52	Bladder, urinary		Normal	E6	F	47	Cervix	Normal	Normal	
A7	M	53	Bladder, urinary		Normal	E7	F	46	Cervix	Normal	Normal	
A8	F	45	Bone, bone marrow		Normal	E8	F	40	Cervix	Normal	Normal	
A9	M	44	Head and neck, salivary glands		Normal	E9	M	42	Pancreas	Normal	Normal	
A10	M	76	Head and neck, salivary glands		Normal	E10	F	26	Pancreas	Normal	Normal	
A11	M	35	Eye	Normal	Normal	E11	F	39	Pancreas	Normal	Normal	
A12	M	50	Eye		Normal	E12	M	77	Peritonsillar	Adenoma	Adenoma	
A13	F	38	Breast		Normal	E13	M	72	Peritonsillar	Adenoma	Adenoma	
B1	F	35	Breast		Normal	F1	F	3	Pituitary gland	Normal	Normal	
B2	F	33	Breast		Normal	F2	F	32	Pituitary gland	Normal	Normal	
B3	M	56	Brain, cerebellum		Normal	F3	F	30	Pituitary gland	Normal	Normal	
B4	M	65	Brain, cerebellum		Normal	F4	F	30	Pituitary gland	Normal	Normal	
B5	F	58	Brain, cerebellum		Normal	F5	F	27	Pituitary gland	Normal	Normal	
B6	M	58	Brain, cerebral cortex		Normal	F6	M	94	Prostate	Normal	Normal	
B7	M	55	Brain, cerebral cortex		Normal	F7	M	95	Prostate	Normal	Normal	
B8	F	42	Fallopian tube	Normal	Normal	F8	M	65	Prostate	Normal	Normal	
B9	F	42	Fallopian tube		Normal	F9	F	32	Skin	Normal	Normal	
B10	F	24	Fallopian tube		Normal	F10	F	86	Skin	Normal	Normal	
B11	F	24	Fallopian tube		Normal	F11	M	26	Skin	Normal	Normal	
B12	M	45	Esophagus		Normal	F12	M	47	Spinal cord	Normal	Normal	
B13	F	66	Esophagus		Normal	F13	M	36	Spinal cord	Normal	Normal	
C1	M	54	Esophagus		Normal	G1	M	27	Spleen	Normal	Normal	
C2	M	45	Stomach		Normal	G2	M	90	Spleen	Normal	Normal	
C3	M	49	Stomach		Normal	G3	M	31	Spleen	Normal	Normal	
C4	M	77	Stomach		Normal	G4	F	90	Skeletal muscle	Normal	Normal	
C5	M	45	Intestine, small intestine	Normal	Normal	G5	F	49	Skeletal muscle	Normal	Normal	
C6	F	75	Intestine, small intestine		Normal	G6	M	70	Skeletal muscle	Normal	Normal	
C7	M	2	Intestine, small intestine		Normal	G7	M	43	Testis	Normal	Normal	
C8	M	46	Intestine, colon		Normal	G8	M	30	Testis	Normal	Normal	
C9	M	46	Intestine, colon		Normal	G9	M	77	Testis	Normal	Normal	
C10	M	74	Intestine, colon		Normal	G10	M	15	Thymus	Normal	Normal	
C11	F	47	Intestine, rectum		Normal	G11	M	26	Thymus	Normal	Normal	
C12	F	75	Intestine, rectum		Normal	G12	F	9	Thymus	Normal	Normal	
C13	M	96	Intestine, rectum		Normal	G13	-	-	-	-	-	-
D1	M	56	Heart		Normal	Normal	H1	F	26	Thyroid	Normal	Normal
D2	M	46	Heart	Normal		H2	F	37	Thyroid	Normal	Normal	
D3	F	34	Heart	Normal		H3	F	51	Thyroid	Normal	Normal	
D4	F	52	Kidney, cortex	Normal		H4	M	46	Tonsil	Normal	Normal	
D5	M	23	Kidney, cortex	Normal		H5	M	37	Tonsil	Normal	Normal	
D6	M	63	Kidney, cortex	Normal		H6	M	13	Tonsil	Normal	Normal	
D7	F	52	Kidney, medulla	Normal		H7	F	36	Uterus, cervix	Normal	Normal	
D8	M	23	Kidney, medulla	Normal		H8	F	35	Uterus, cervix	Normal	Normal	
D9	M	59	Peripheral nerve	Schwannoma		H9	F	44	Uterus, cervix	Normal	Normal	
D10	F	29	Peripheral nerve	Schwannoma		H10	F	38	Uterus, endometrium	Normal	Normal	
D11	F	44	Ureter	Normal	Normal	H11	F	41	Uterus, endometrium	Normal	Normal	
D12	F	42	Ureter		Normal	H12	F	48	Uterus, endometrium	Normal	Normal	
D13	F	57	Liver		Normal	H13	M	58	Skin	Malignant melanoma (tissue marker)	Malignant melanoma (tissue marker)	

Fig. 153B

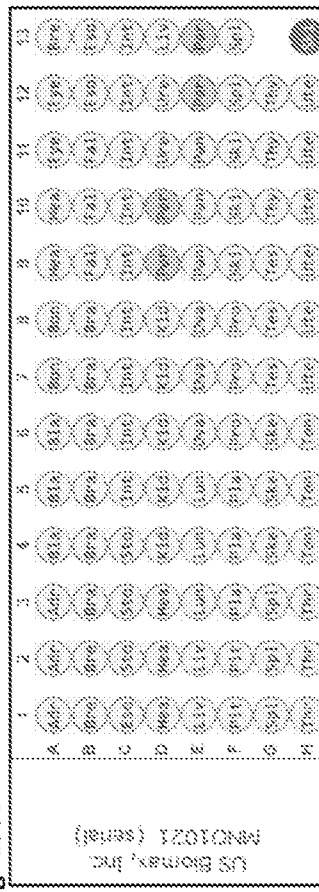


Figure 153A-153C

PSMGFR-antibody 28F9 IgG 15 ug/mL  
FDA Normal tissue array MNO1021

Fig. 154A

Normal Adrenal Gland

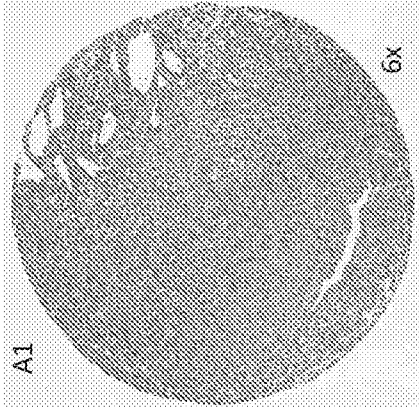


Fig. 154B

Normal Breast

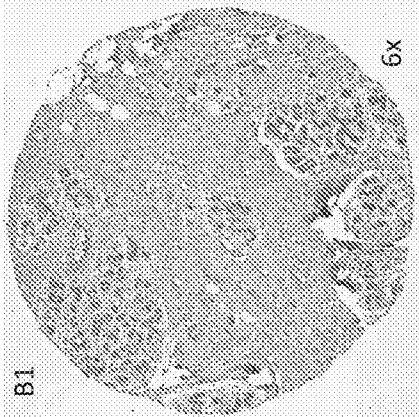


Fig. 154C

Normal Fallopian Tubes

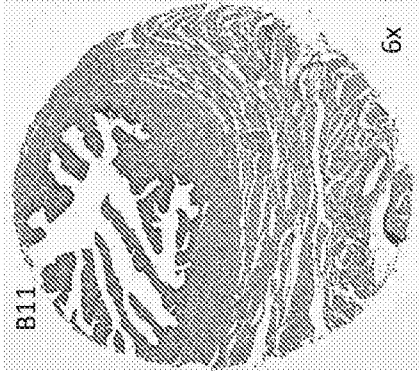


Fig. 154D

Normal Kidney

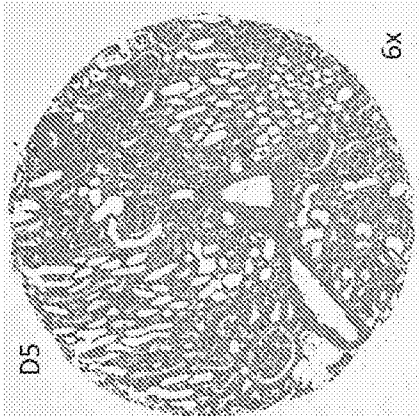


Fig. 154E

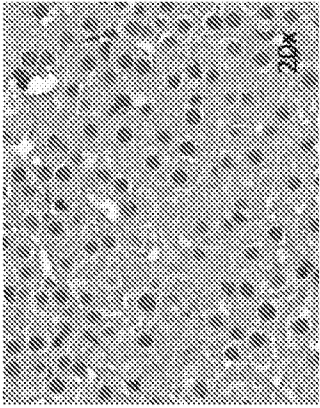


Fig. 154F



Fig. 154G

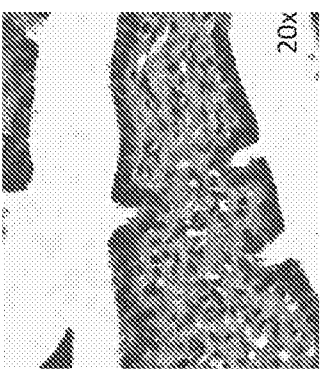


Fig. 154H

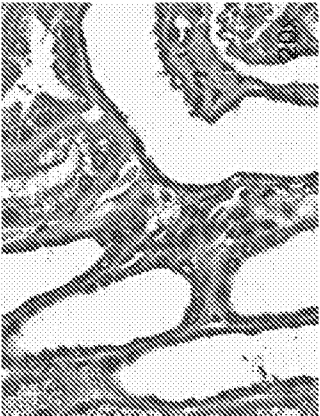


Figure 154A-154H

PSMGFR-antibody 28F9 IgG 15 ug/mL  
FDA Normal tissue array MNO1021

Fig. 154I  
Normal Heart

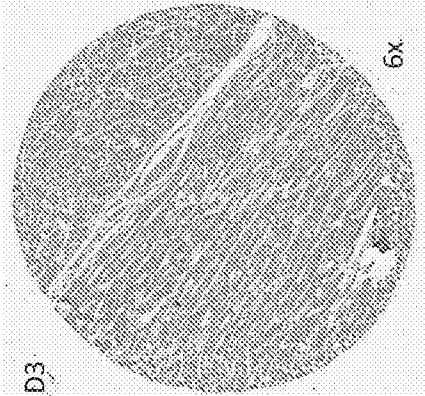


Fig. 154J  
Normal Liver

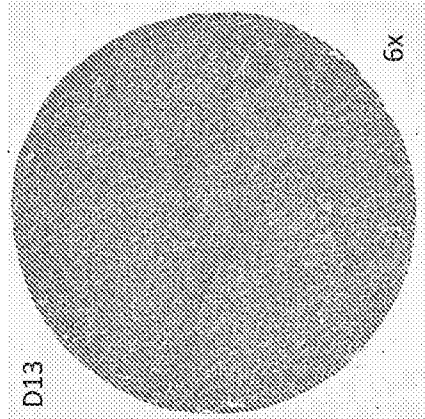


Fig. 154K  
Normal Lung

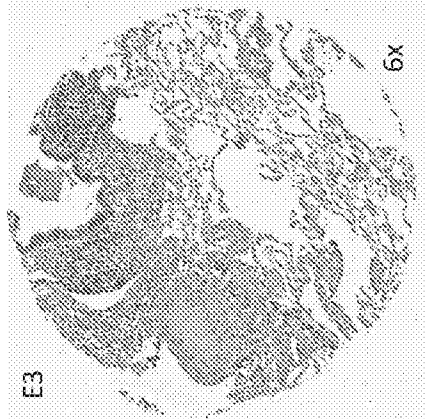


Fig. 154L  
Normal Ureter

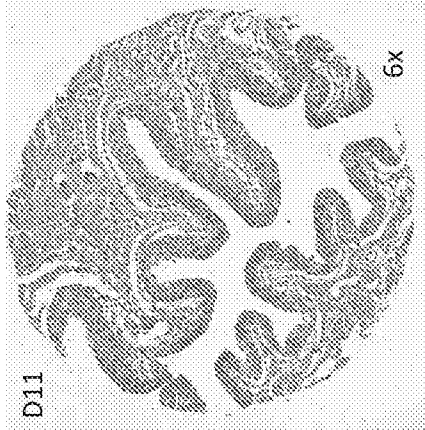


Fig. 154M

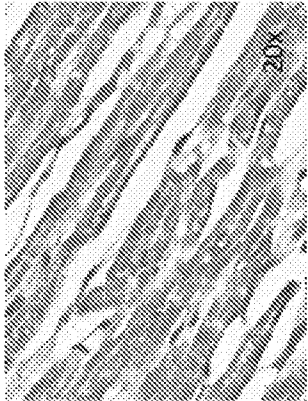


Fig. 154N

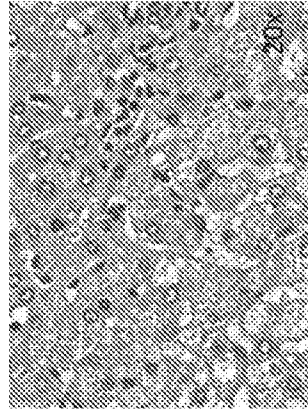


Fig. 154O



Fig. 154P

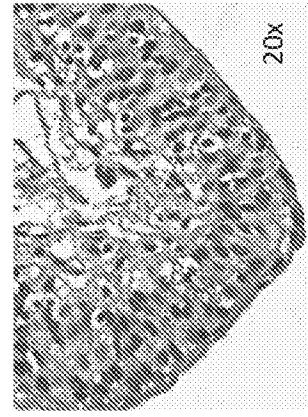


Figure 154I-154P

PSMGFR-antibody 28F9 IgG 15 ug/mL  
FDA Normal tissue array MNO1021

Fig. 154Q  
Normal eye

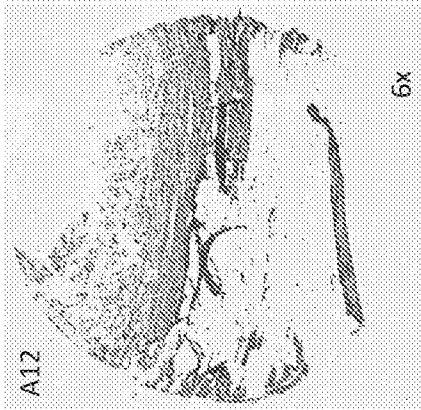


Fig. 154R  
Normal Cerebral cortex

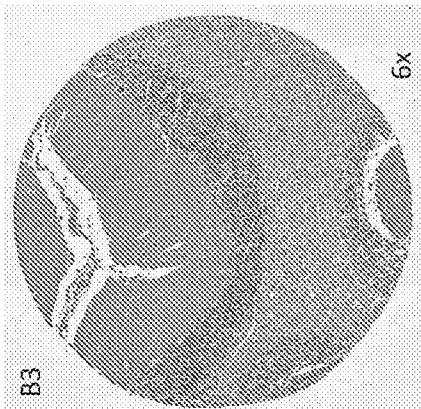


Fig. 154S  
Normal Bone marrow

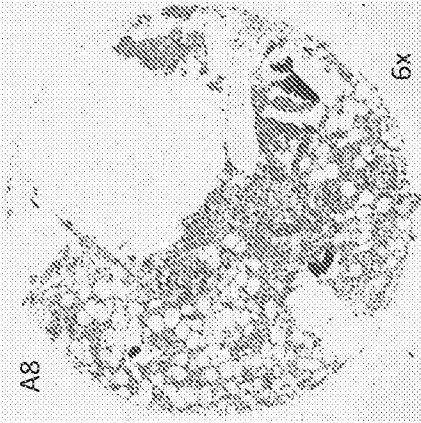


Fig. 154T  
Normal Skeletal muscle

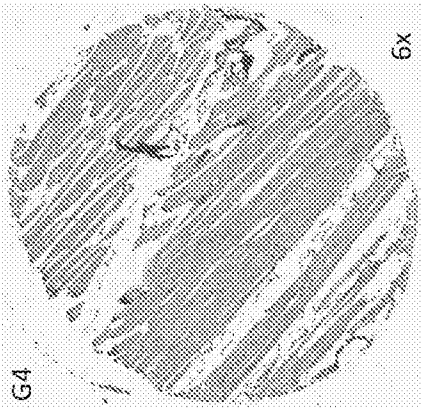


Fig. 154U



Fig. 154V

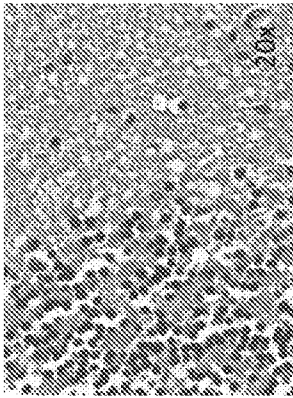


Fig. 154W

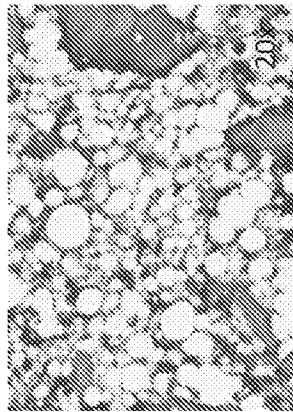


Fig. 154X

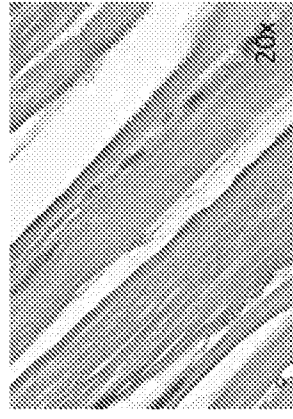


Figure 154Q-154X

100-1-1-1 Form 101

100-1-1-1 Form 101

[illegible]

Figure 15A-15C

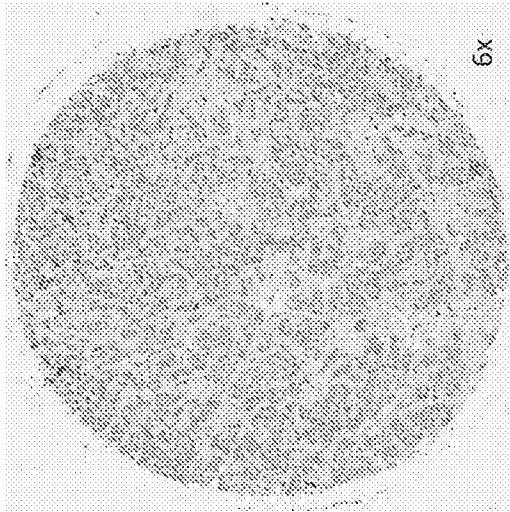
[illegible]



**PSMGFR-antibody 28F9 15ug/mL**  
Breast cancer tissue array BR1141

**Fig. 156A**

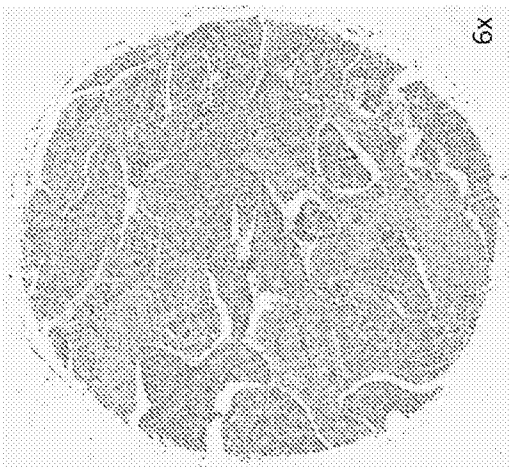
Position: C5  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



6x

**Fig. 156B**

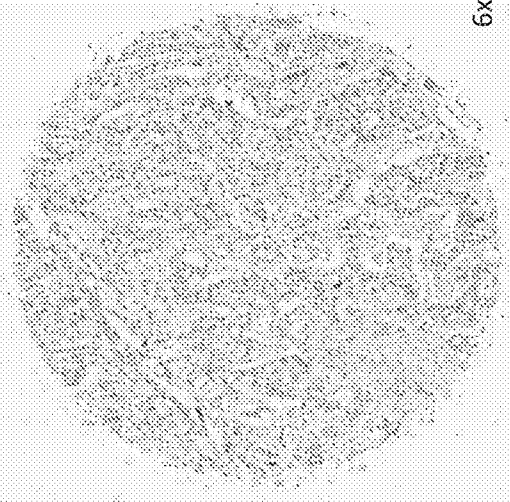
Position: G9  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T1N0M0



6x

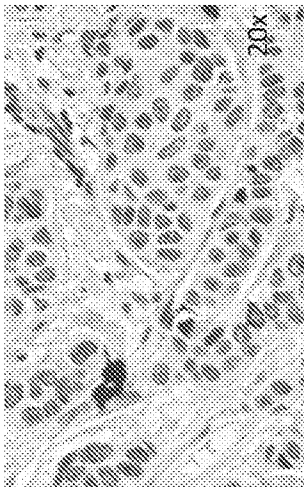
**Fig. 156C**

Position: I1  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N1M0



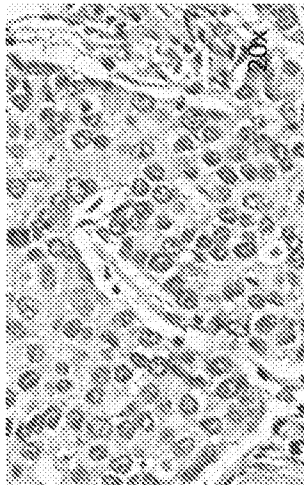
6x

**Fig. 156D**



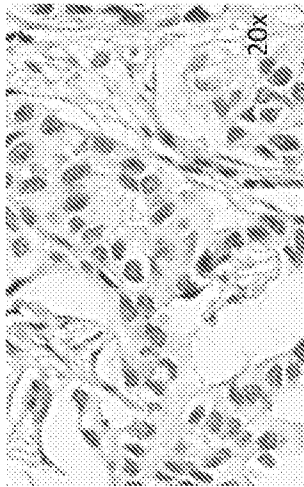
20x

**Fig. 156E**



20x

**Fig. 156F**



20x

**Fig. 156A-156F**

N+20/C-27 antibody 1E4 IgG 7.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 157A

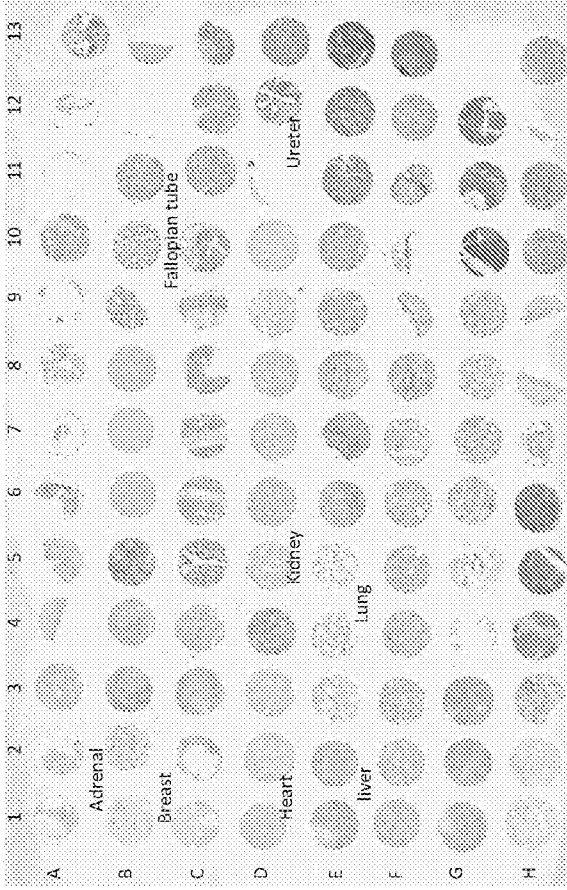


Fig. 157C

Position	Age	Sex	Organ/Anatomic Site	Pathology	Diagnosis	Position	Age	Sex	Organ/Anatomic Site	Pathology	Diagnosis
A1	F	31	Adrenal gland	Normal	Normal	E1	M	59	Liver	Normal	Normal
A2	M	51	Adrenal gland	Normal	Normal	E2	F	27	Liver	Normal	Normal
A3	M	51	Adrenal gland	Normal	Normal	E3	M	72	Lung	Normal	Normal
A4	M	72	Bladder, urinary	Normal	Normal	E4	F	55	Lung	Normal	Normal
A5	M	72	Bladder, urinary	Normal	Normal	E5	M	45	Lung	Normal	Normal
A6	M	52	Bladder, urinary	Normal	Normal	E6	F	47	Ovary	Normal	Normal
A7	M	52	Bone, bone marrow	Normal	Normal	E7	F	46	Ovary	Normal	Normal
A8	F	43	Bone, bone marrow	Normal	Normal	E8	F	46	Ovary	Normal	Normal
A9	F	44	Head and neck, salivary gland	Normal	Normal	E9	M	42	Parotids	Normal	Normal
A10	M	18	Head and neck, salivary gland	Normal	Normal	E10	F	26	Parotids	Normal	Normal
A11	M	55	Eye	Normal	Normal	E11	M	59	Parotids	Normal	Normal
A12	M	50	Eye	Normal	Normal	E12	M	17	Parathyroid	Adenoma	Adenoma
A13	F	35	Breast	Normal	Normal	E13	F	72	Parathyroid	Adenoma	Adenoma
B1	F	38	Breast	Normal	Normal	F1	F	3	Mammary gland	Normal	Normal
B2	F	38	Breast	Normal	Normal	F2	F	32	Mammary gland	Normal	Normal
B3	M	58	Brain, cerebellum	Normal	Normal	F3	F	30	Placenta	Normal	Normal
B4	M	58	Brain, cerebellum	Normal	Normal	F4	F	27	Placenta	Normal	Normal
B5	M	58	Brain, cerebral cortex	Normal	Normal	F5	M	64	Prostate	Normal	Normal
B6	M	58	Brain, cerebral cortex	Normal	Normal	F6	M	65	Prostate	Normal	Normal
B7	F	42	Fallopian tube	Normal	Normal	F7	M	65	Prostate	Normal	Normal
B8	F	42	Fallopian tube	Normal	Normal	F8	M	65	Prostate	Normal	Normal
B9	F	32	Fallopian tube	Normal	Normal	F9	F	32	Skin	Normal	Normal
B10	F	32	Fallopian tube	Normal	Normal	F10	F	26	Skin	Normal	Normal
B11	F	32	Fallopian tube	Normal	Normal	F11	M	26	Skin	Normal	Normal
B12	F	34	Esophagus	Normal	Normal	F12	M	58	Spinal cord	Normal	Normal
B13	F	34	Esophagus	Normal	Normal	G1	M	27	Spleen	Normal	Normal
C1	M	45	Esophagus	Normal	Normal	G2	M	30	Spleen	Normal	Normal
C2	M	45	Esophagus	Normal	Normal	G3	M	31	Spleen	Normal	Normal
C3	M	45	Esophagus	Normal	Normal	G4	F	60	Skeletal muscle	Normal	Normal
C4	M	45	Esophagus	Normal	Normal	G5	M	49	Skeletal muscle	Normal	Normal
C5	M	45	Esophagus	Normal	Normal	G6	M	70	Skeletal muscle	Normal	Normal
C6	F	75	Intestine, small intestine	Normal	Normal	G7	M	43	Testis	Normal	Normal
C7	F	75	Intestine, small intestine	Normal	Normal	G8	M	40	Testis	Normal	Normal
C8	M	45	Intestine, small intestine	Normal	Normal	G9	M	43	Testis	Normal	Normal
C9	M	45	Intestine, small intestine	Normal	Normal	G10	M	15	Thymus	Normal	Normal
C10	M	45	Intestine, small intestine	Normal	Normal	G11	M	26	Thymus	Normal	Normal
C11	F	75	Intestine, colon	Normal	Normal	G12	F	9	Thymus	Normal	Normal
C12	F	75	Intestine, colon	Normal	Normal	G13	F	26	Thyroid	Normal	Normal
C13	M	36	Intestine, rectum	Normal	Normal	H1	F	37	Thyroid	Normal	Normal
D1	M	36	Intestine, rectum	Normal	Normal	H2	F	51	Thyroid	Normal	Normal
D2	M	36	Intestine, rectum	Normal	Normal	H3	F	51	Thyroid	Normal	Normal
D3	F	34	Heart	Normal	Normal	H4	M	46	Tonsil	Normal	Normal
D4	F	34	Heart	Normal	Normal	H5	M	37	Tonsil	Normal	Normal
D5	F	52	Kidney, cortex	Normal	Normal	H6	M	13	Tonsil	Normal	Normal
D6	F	52	Kidney, cortex	Normal	Normal	H7	F	36	Uterus, cervix	Normal	Normal
D7	M	53	Kidney, medulla	Normal	Normal	H8	F	35	Uterus, cervix	Normal	Normal
D8	M	53	Kidney, medulla	Normal	Normal	H9	F	44	Uterus, cervix	Normal	Normal
D9	M	53	Kidney, medulla	Normal	Normal	H10	F	36	Uterus	Normal	Normal
D10	F	28	Peripheral nerve	Schwannoma	Schwannoma	H11	F	41	Uterus	Normal	Normal
D11	F	44	Peripheral nerve	Schwannoma	Schwannoma	H12	F	46	Uterus	Normal	Normal
D12	F	44	Ureter	Normal	Normal	H13	M	56	Skin	Malignant melanoma (tissue marker)	Malignant melanoma (tissue marker)
D13	F	44	Ureter	Normal	Normal						

Fig. 157B

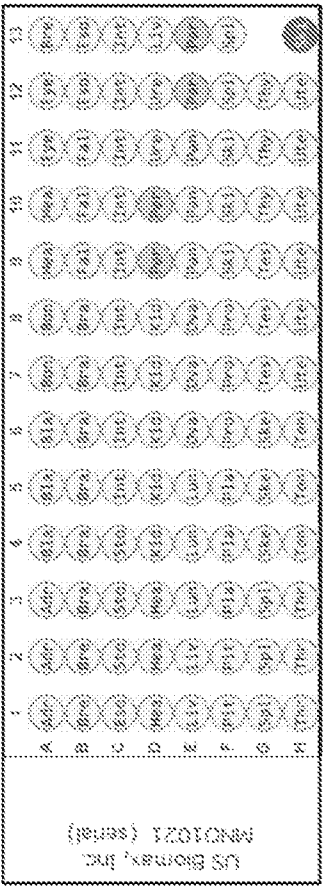


Figure 157A-157C



N+20/C-27 antibody 1E4 IgG 7.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 158A

Normal Adrenal Gland

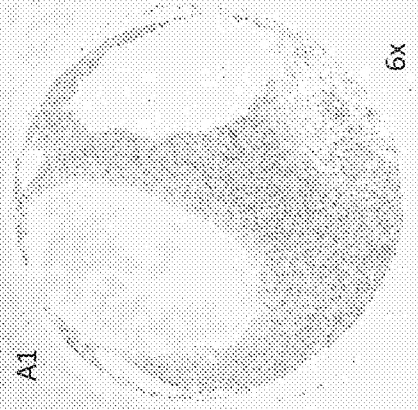


Fig. 158B

Normal Breast

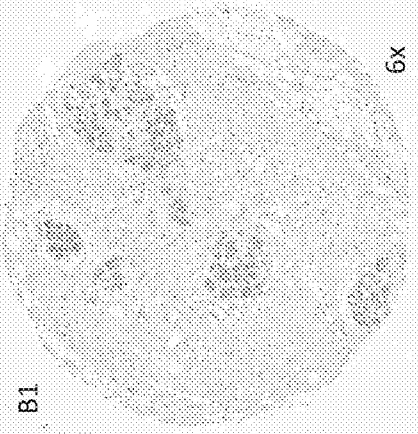


Fig. 158C

Normal Fallopian Tubes

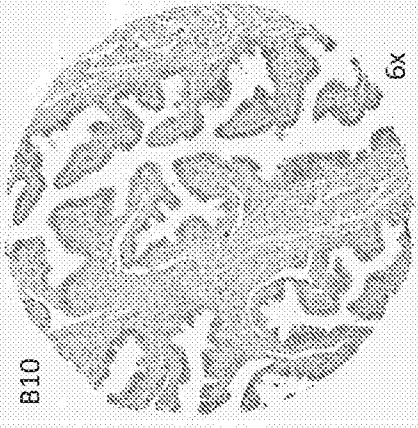


Fig. 158D

Normal Kidney

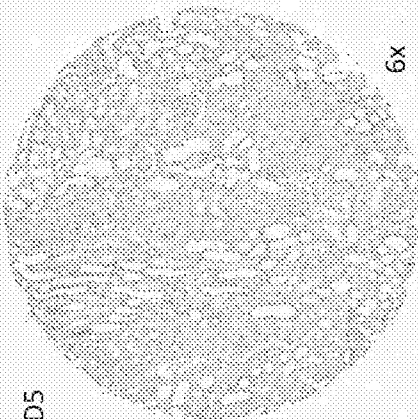


Fig. 158E

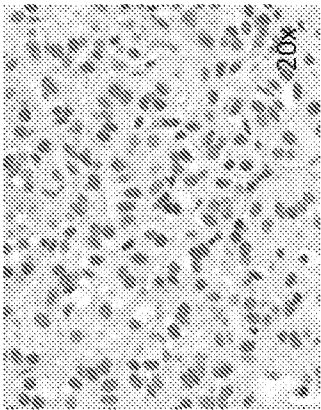


Fig. 158F

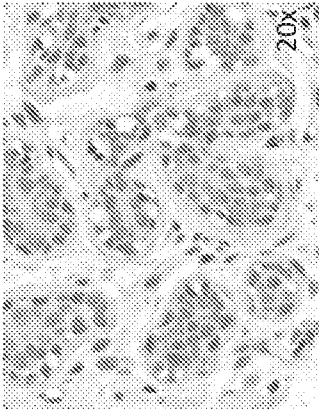


Fig. 158G

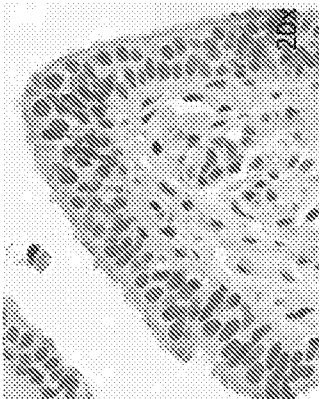


Fig. 158H

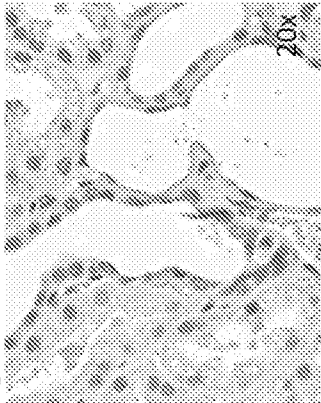


Figure 158A-158H

N+20/C-27 antibody 1E4 IgG 7.5  
ug/mL  
FDA Normal tissue array MNO1021

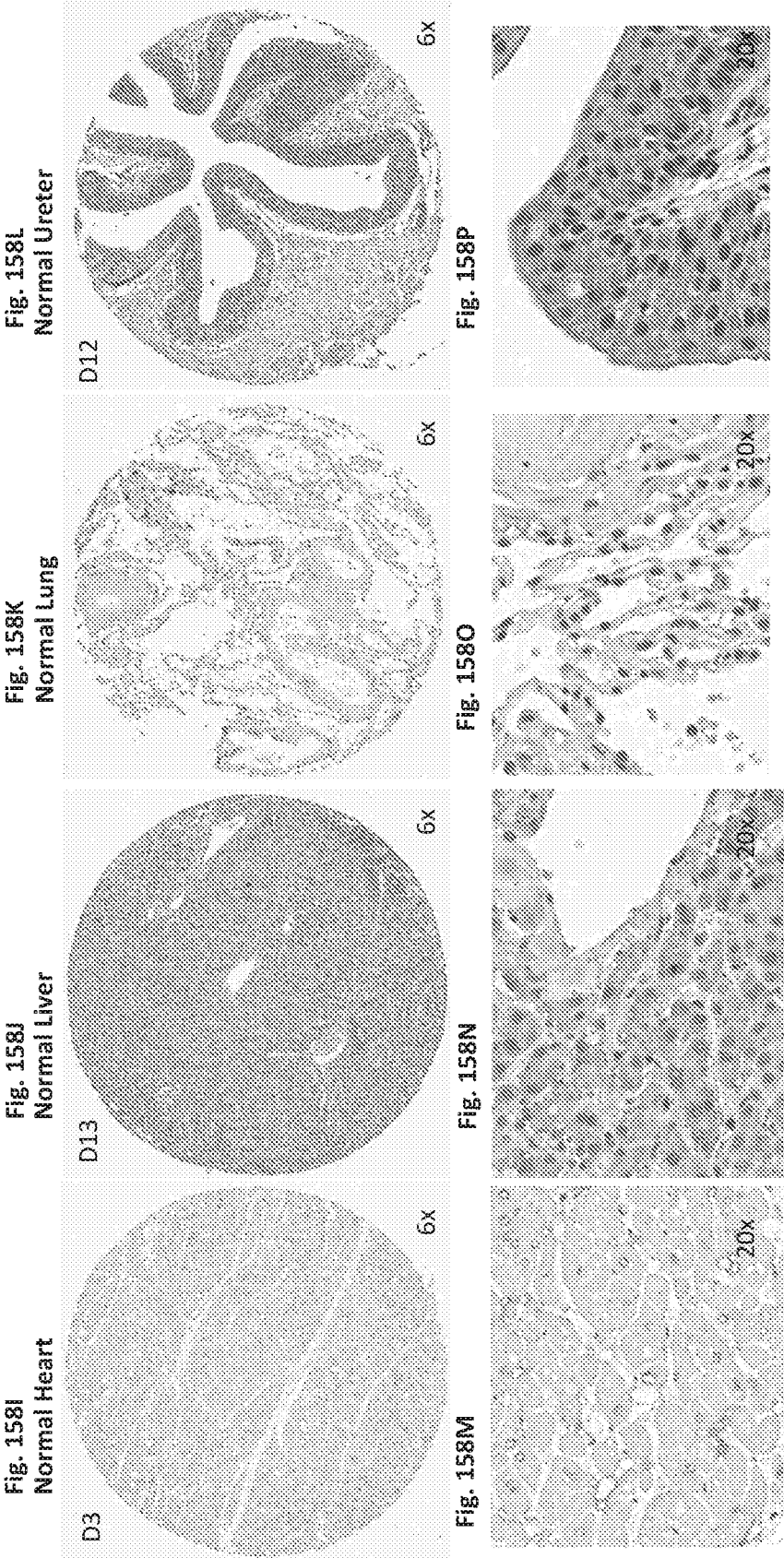


Figure 158I-158P

N+20/C-27 antibody 1E4 IgG 7.5  
ug/mL  
FDA Normal tissue array MNO1021

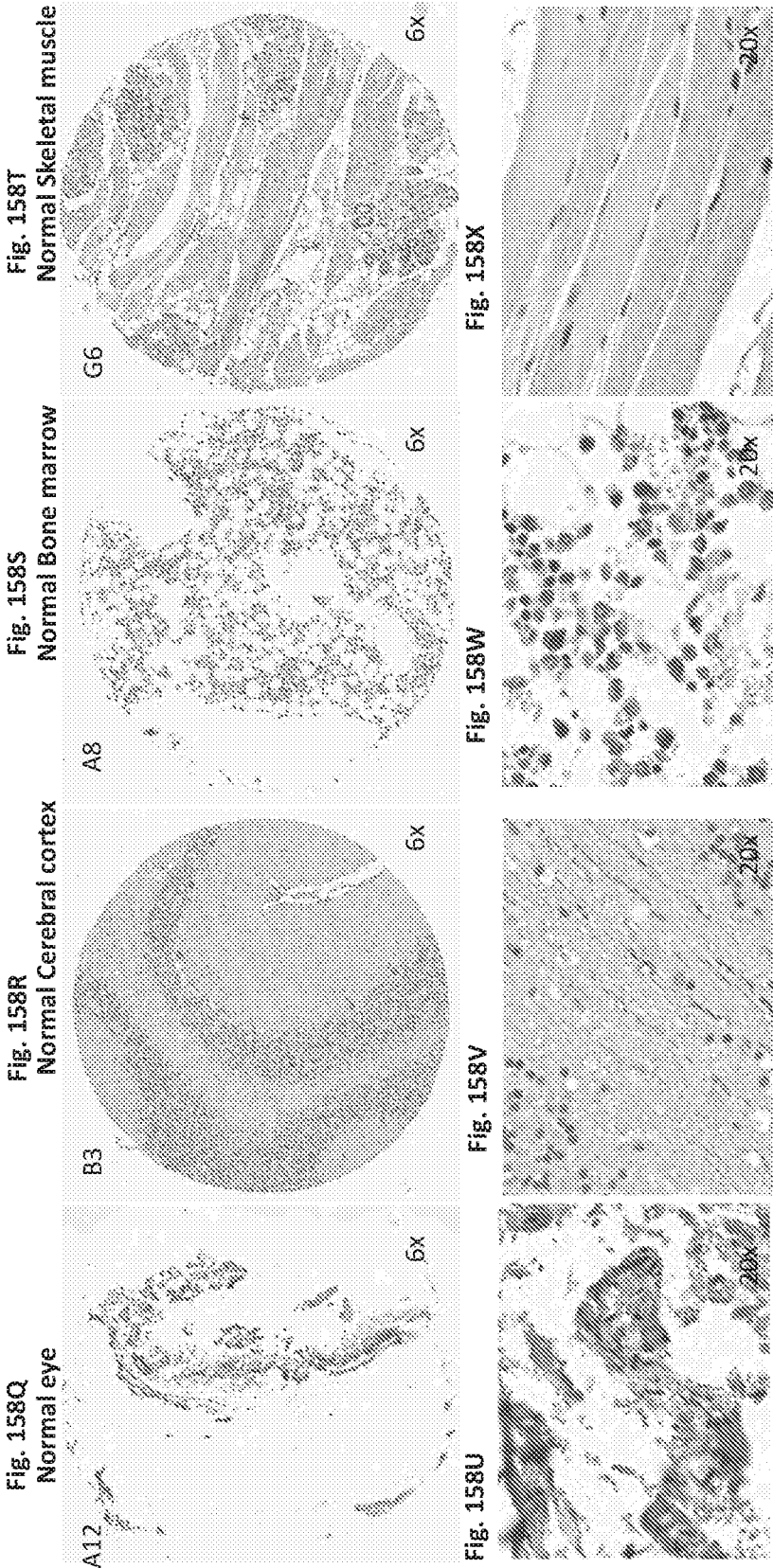
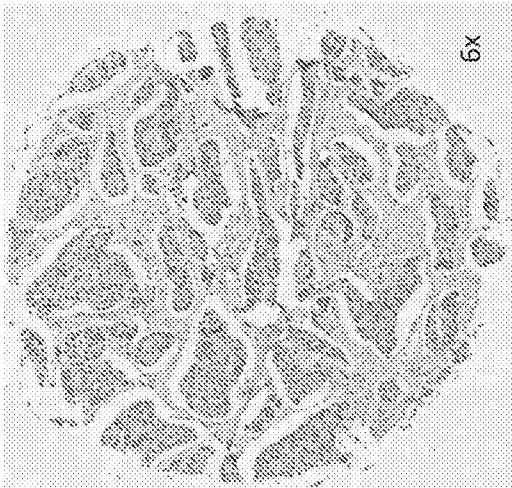


Figure 158Q-158X

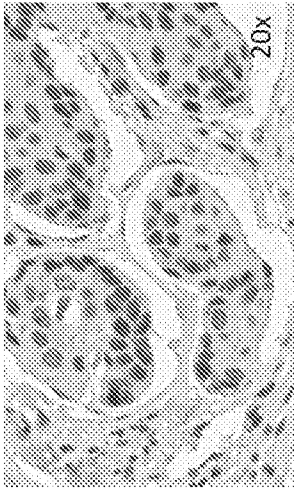


N+20/C-27 antibody 1E4 IgG 10 ug/mL  
Breast cancer tissue array BR1007

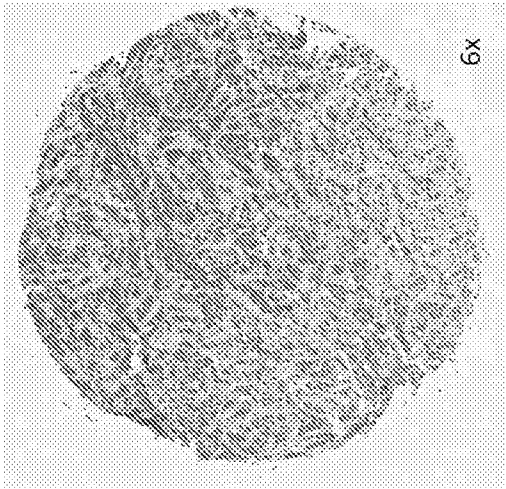
**Fig. 160A**  
Position: B4  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T1N1M0



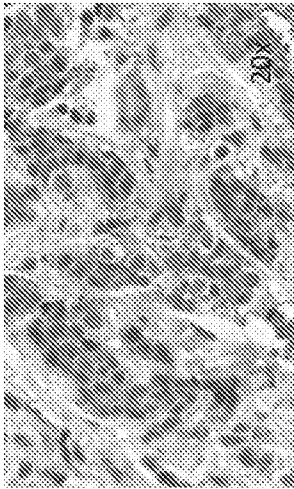
**Fig. 160D**



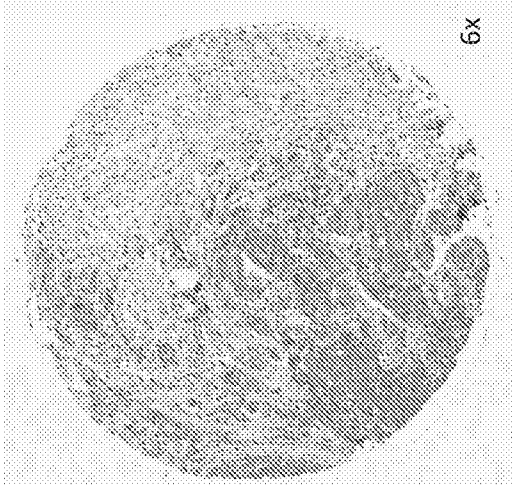
**Fig. 160B**  
Position: C4  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T4N0M0



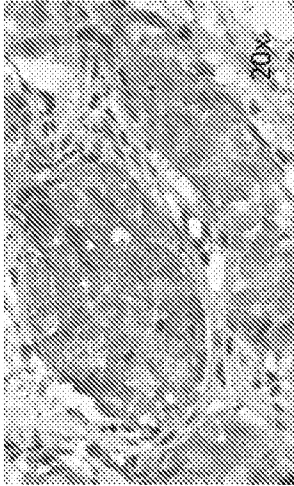
**Fig. 160E**



**Fig. 160C**  
Position: D5  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



**Fig. 160F**



**Fig. 160A-- 160F**

N+20/C-27 antibody 29H1 IgG 0.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 161A

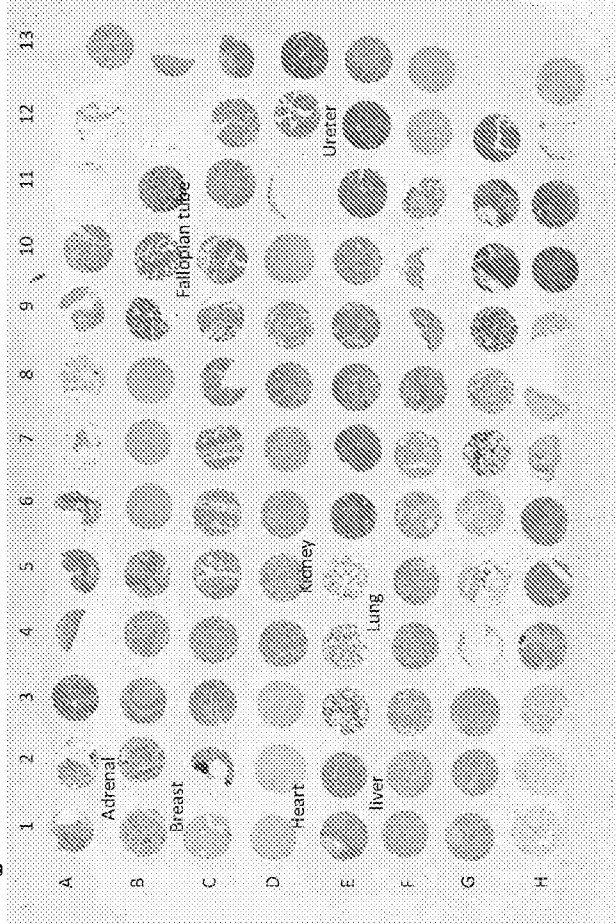


Fig. 161C

Position	Age	Sex	Organ/Anatomic Site	Pathology	Diagnosis	Position	Age	Sex	Organ/Anatomic Site	Pathology	Diagnosis
A1	F	31	Adrenal gland	Normal	Normal	E1	M	59	Liver	Normal	Normal
A2	M	51	Adrenal gland	Normal	Normal	E2	F	27	Liver	Normal	Normal
A3	M	51	Adrenal gland	Normal	Normal	E3	F	72	Lung	Normal	Normal
A4	M	72	Bladder, urinary	Normal	Normal	E4	M	55	Lung	Normal	Normal
A5	M	52	Bladder, urinary	Normal	Normal	E5	M	45	Lung	Normal	Normal
A6	M	53	Bone, bone marrow	Normal	Normal	E6	F	47	Ovary	Normal	Normal
A7	M	53	Bone, bone marrow	Normal	Normal	E7	F	46	Ovary	Normal	Normal
A8	F	44	Head and neck, salivary gland	Normal	Normal	E8	F	40	Ovary	Normal	Normal
A9	M	18	Head and neck, salivary gland	Normal	Normal	E9	M	42	Parotids	Normal	Normal
A10	M	55	Eye	Normal	Normal	E10	F	26	Parotids	Normal	Normal
A11	M	55	Eye	Normal	Normal	E11	F	59	Parotids	Normal	Normal
A12	M	50	Eye	Normal	Normal	E12	M	17	Parathyroid	Adenoma	Adenoma
A13	F	35	Breast	Normal	Normal	E13	F	72	Parathyroid	Adenoma	Adenoma
B1	F	38	Breast	Normal	Normal	F1	F	3	Mammary gland	Normal	Normal
B2	F	38	Breast	Normal	Normal	F2	F	32	Mammary gland	Normal	Normal
B3	M	58	Brain, cerebellum	Normal	Normal	F3	F	30	Mammary gland	Normal	Normal
B4	M	58	Brain, cerebellum	Normal	Normal	F4	F	27	Mammary gland	Normal	Normal
B5	M	58	Brain, cerebral cortex	Normal	Normal	F5	M	64	Prostate	Normal	Normal
B6	M	58	Brain, cerebral cortex	Normal	Normal	F6	M	65	Prostate	Normal	Normal
B7	F	42	Fallopian tube	Normal	Normal	F7	M	65	Prostate	Normal	Normal
B8	F	32	Fallopian tube	Normal	Normal	F8	M	32	Skin	Normal	Normal
B9	F	32	Fallopian tube	Normal	Normal	F9	F	26	Skin	Normal	Normal
B10	F	24	Fallopian tube	Normal	Normal	F10	M	26	Skin	Normal	Normal
B11	F	24	Fallopian tube	Normal	Normal	F11	M	26	Skin	Normal	Normal
B12	M	34	Esophagus	Normal	Normal	F12	M	58	Spinal cord	Normal	Normal
B13	M	34	Esophagus	Normal	Normal	F13	M	27	Spinal cord	Normal	Normal
C1	M	45	Stomach	Normal	Normal	G1	M	27	Spleen	Normal	Normal
C2	M	45	Stomach	Normal	Normal	G2	M	30	Spleen	Normal	Normal
C3	M	77	Stomach	Normal	Normal	G3	M	31	Spleen	Normal	Normal
C4	M	77	Stomach	Normal	Normal	G4	F	60	Skeletal muscle	Normal	Normal
C5	M	45	Intestine, small intestine	Normal	Normal	G5	M	49	Skeletal muscle	Normal	Normal
C6	F	75	Intestine, small intestine	Normal	Normal	G6	M	70	Skeletal muscle	Normal	Normal
C7	F	75	Intestine, small intestine	Normal	Normal	G7	M	43	Testis	Normal	Normal
C8	M	46	Intestine, colon	Normal	Normal	G8	M	40	Testis	Normal	Normal
C9	M	46	Intestine, colon	Normal	Normal	G9	M	77	Testis	Normal	Normal
C10	M	74	Intestine, colon	Normal	Normal	G10	M	15	Thymus	Normal	Normal
C11	F	77	Intestine, rectum	Normal	Normal	G11	M	26	Thymus	Normal	Normal
C12	F	77	Intestine, rectum	Normal	Normal	G12	F	9	Thymus	Normal	Normal
C13	M	36	Intestine, rectum	Normal	Normal	G13	F	26	Thyroid	Normal	Normal
D1	M	36	Heart	Normal	Normal	H1	F	26	Thyroid	Normal	Normal
D2	M	36	Heart	Normal	Normal	H2	F	37	Thyroid	Normal	Normal
D3	F	44	Heart	Normal	Normal	H3	F	51	Thyroid	Normal	Normal
D4	F	52	Kidney, cortex	Normal	Normal	H4	M	46	Tonsil	Normal	Normal
D5	M	23	Kidney, cortex	Normal	Normal	H5	M	37	Tonsil	Normal	Normal
D6	M	53	Kidney, medulla	Normal	Normal	H6	M	13	Tonsil	Normal	Normal
D7	M	53	Kidney, medulla	Normal	Normal	H7	F	36	Uterus, cervix	Normal	Normal
D8	M	53	Kidney, medulla	Normal	Normal	H8	F	35	Uterus, cervix	Normal	Normal
D9	M	39	Peripheral nerve	Schwannoma	Schwannoma	H9	F	44	Uterus, cervix	Normal	Normal
D10	F	28	Peripheral nerve	Schwannoma	Schwannoma	H10	F	36	Uterus	Normal	Normal
D11	F	44	Ureter	Normal	Normal	H11	F	41	Uterus	Normal	Normal
D12	F	42	Ureter	Normal	Normal	H12	F	46	Uterus	Normal	Normal
D13	F	57	Liver	Normal	Normal	H13	M	56	Skin	Malignant melanoma (tissue marker)	Malignant melanoma (tissue marker)

Fig. 161B

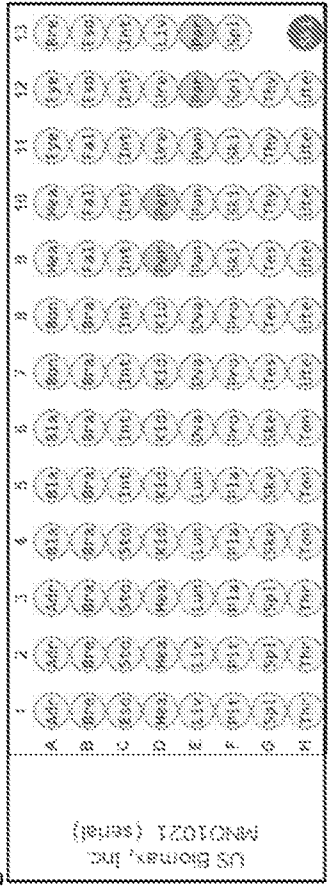


Figure 161A-161C



N+20/C-27 antibody 29H1 IgG 0.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 162A

Normal Adrenal Gland

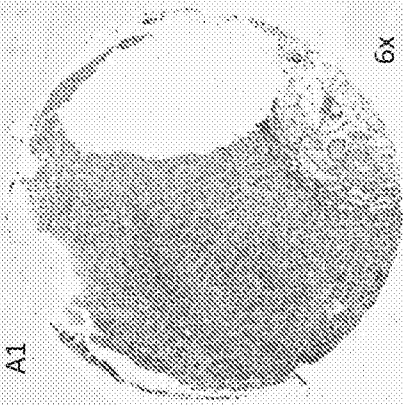


Fig. 162B

Normal Breast

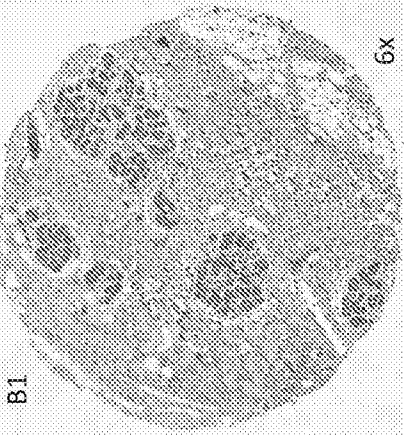


Fig. 162C

Normal Fallopian Tubes

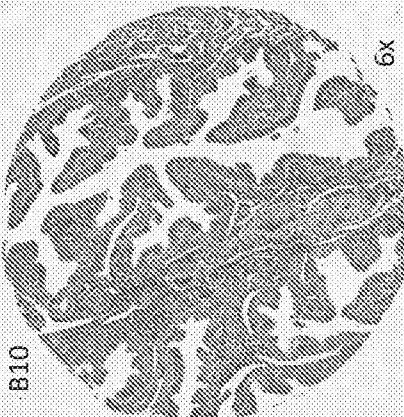


Fig. 162D

Normal Kidney

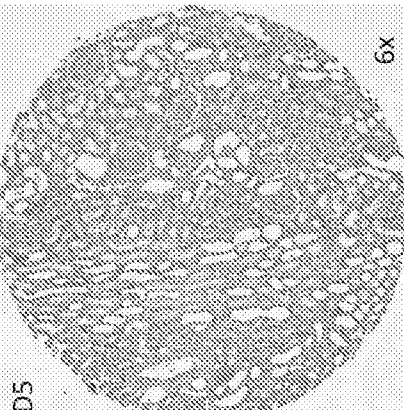


Fig. 162E

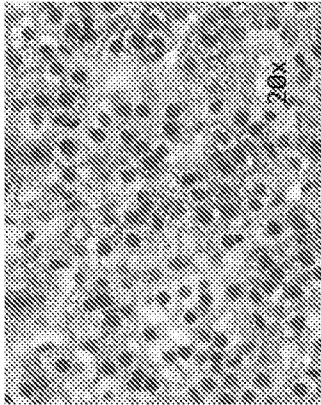


Fig. 162F



Fig. 162G

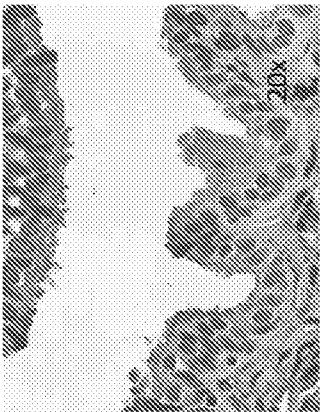


Fig. 162H

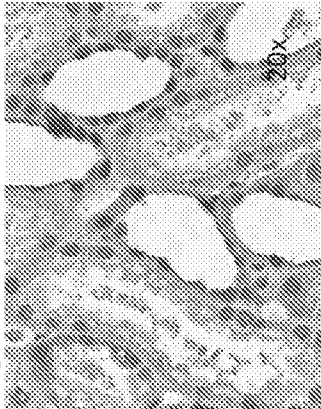


Figure 162A-162H

N+20/C-27 antibody 29H1 IgG 0.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 162I  
Normal Heart

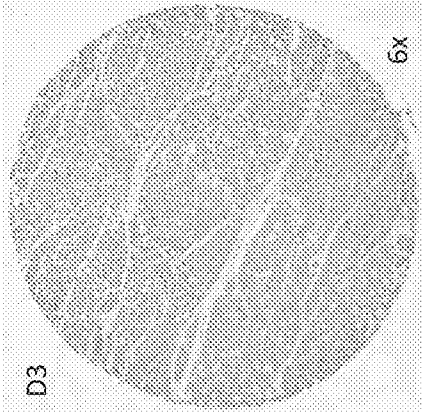


Fig. 162J  
Normal Liver

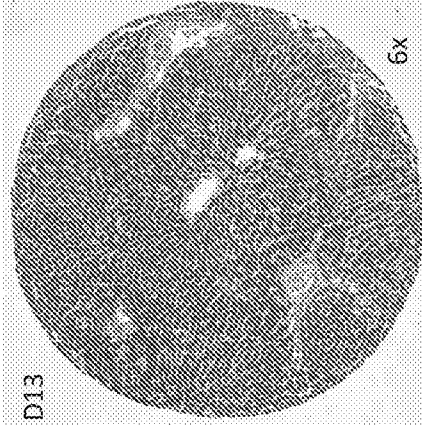


Fig. 162K  
Normal Lung



Fig. 162L  
Normal Ureter

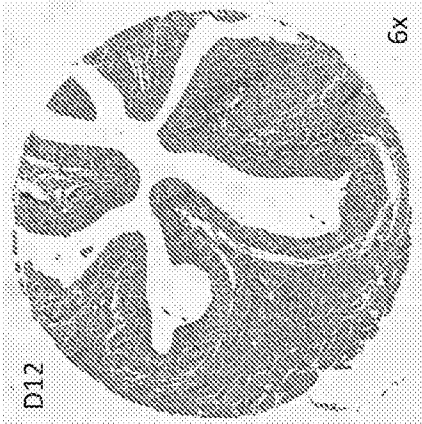


Fig. 162M

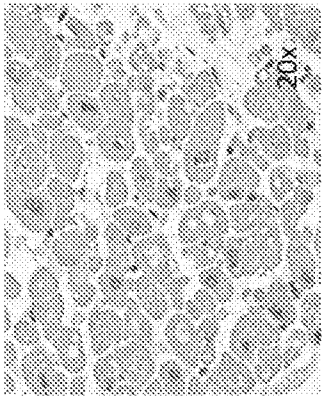


Fig. 162N

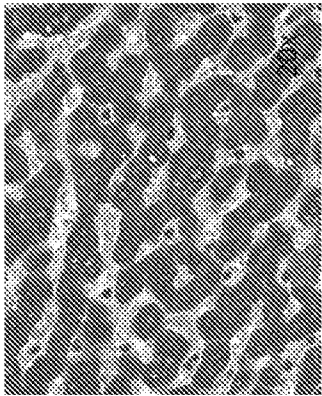


Fig. 162O

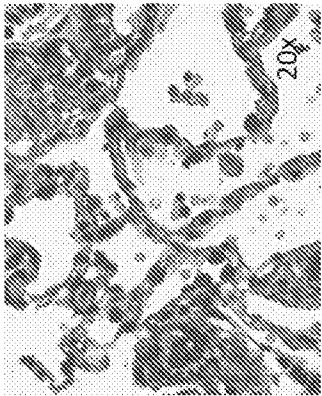


Fig. 162P

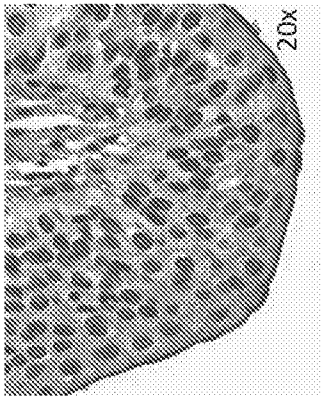


Figure 162I-162P



N+20/C-27 antibody 29H1 IgG 0.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 162Q  
Normal eye

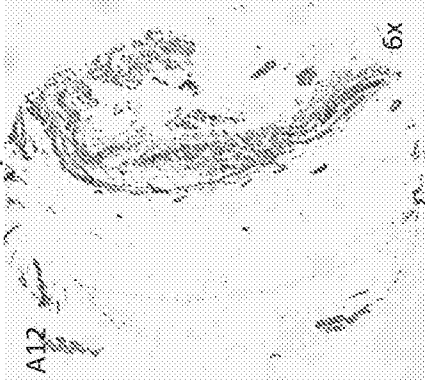


Fig. 162R  
Normal Cerebral cortex

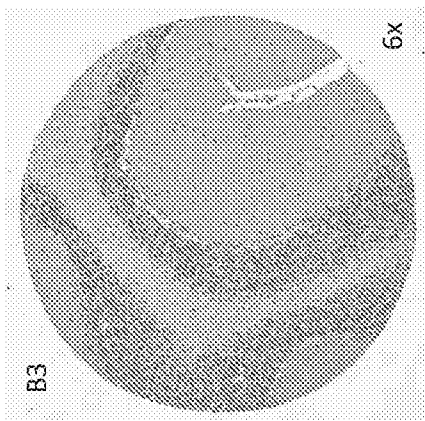


Fig. 162S  
Normal Bone marrow

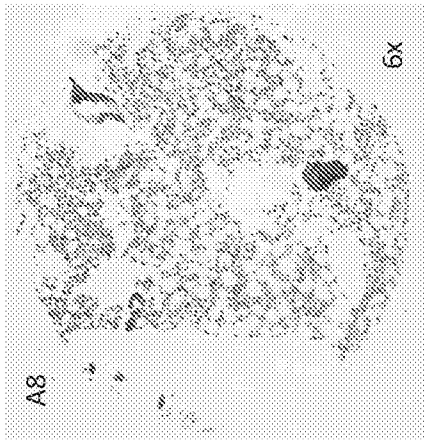


Fig. 162T  
Normal Skeletal muscle

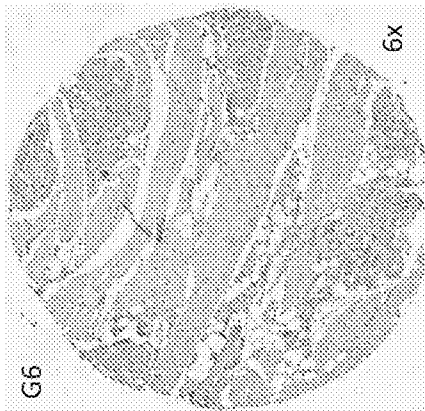


Fig. 162U

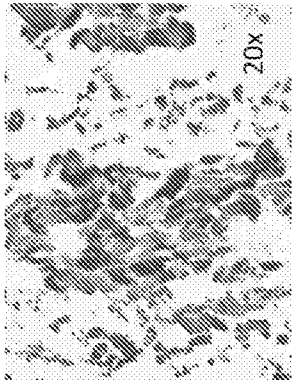


Fig. 162V

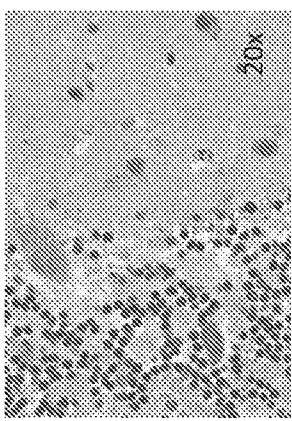


Fig. 162W

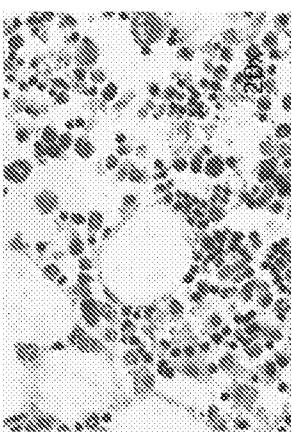


Fig. 162X

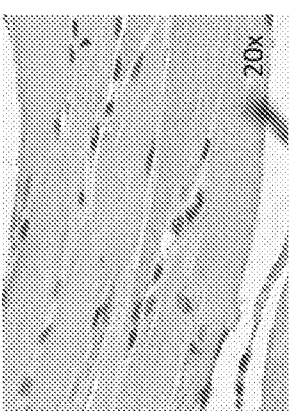


Figure 162Q-162X

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# N+20/C-27 antibody 29H1 IgG 0.5 ug/mL Breast cancer tissue array Br1141

Fig. 163A

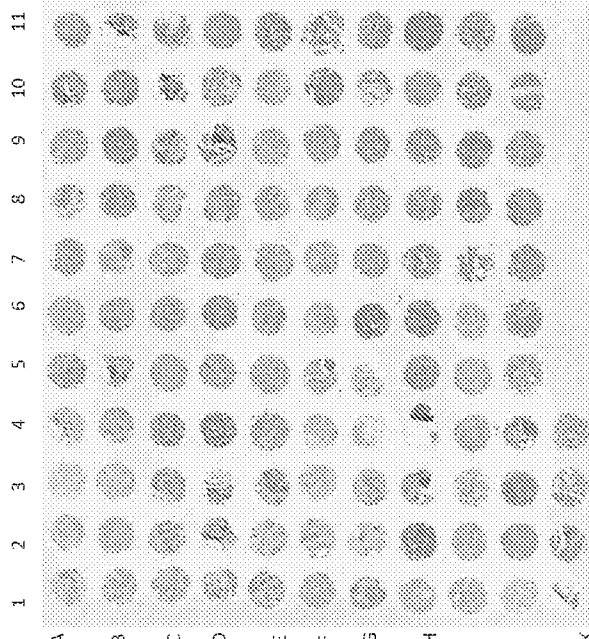


Fig. 163C

Position	Site	Primary diagnosis	TMN	Grade	Type	Position	Site	Primary diagnosis	TMN	Grade	Type				
A1	45	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	F1	52	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
A2	38	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	F2	44	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
A3	42	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	F3	53	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
A4	41	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	F4	94	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
A5	70	F	invasive ductal carcinoma	T2N1M0	2	IA	Malignant	F5	83	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
A6	48	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant	F6	89	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
A7	42	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	F7	43	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
A8	48	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	F8	45	F	invasive ductal carcinoma	T2N1M0	2	IA	Malignant
A9	98	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	F9	82	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
A10	82	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant	F10	42	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
A11	90	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	F11	45	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
B1	43	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	G1	45	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
B2	43	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	G2	42	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
B3	37	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	G3	42	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
B4	77	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	G4	44	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
B5	50	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	G5	45	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
B6	48	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	G6	47	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
B7	48	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	G7	53	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
B8	40	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	G8	42	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
B9	50	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	G9	38	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
B10	47	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	G10	46	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
B11	48	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	H1	45	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
C1	48	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	H2	83	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant
C2	42	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	H3	52	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant
C3	41	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	H4	50	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
C4	95	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	H5	42	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
C5	41	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	H6	42	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
C6	41	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant	H7	47	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
C7	41	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant	H8	47	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
C8	40	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant	H9	45	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
C9	40	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	H10	51	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
C10	42	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	H11	42	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
C11	37	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	I1	45	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
C12	48	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	I2	66	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant
D1	49	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	I3	75	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
D2	48	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	I4	98	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
D3	48	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	I5	24	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
D4	48	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	I6	35	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant
D5	52	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	I7	35	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant
D6	44	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	I8	39	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
D7	43	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	I9	47	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
D8	44	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	I10	57	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
D9	46	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	I11	73	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
E1	44	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	J1	42	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
E2	97	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	J2	86	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
E3	48	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	J3	45	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
E4	44	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant	J4	44	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant
E5	44	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant	J5	48	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant
E6	44	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant	J6	52	F	invasive ductal carcinoma	T2N0M0	2	IB	Malignant
E7	98	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	J7	56	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
E8	98	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant	J8	42	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
E9	98	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	J9	41	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
E10	48	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	J10	53	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
E11	52	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant	J11	57	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
								K1	45	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
								K2	46	F	invasive ductal carcinoma	T2N0M0	2	IA	Malignant
								K3	46	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant
								K4	46	F	invasive ductal carcinoma	T2N1M0	2	IB	Malignant

Fig. 163B

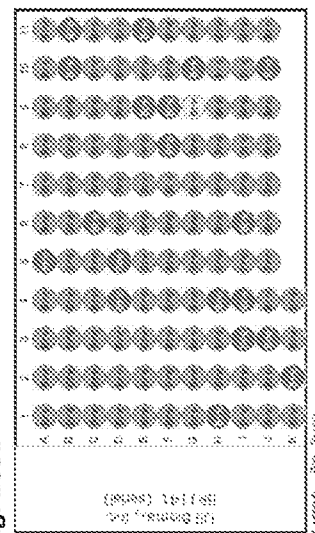
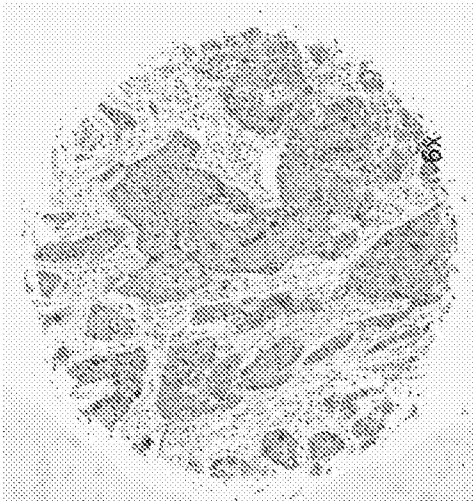


Figure 163A-163C

**N+20/C-27 antibody 29H1 IgG 0.5ug/ml**  
**Breast cancer tissue array Br1141**

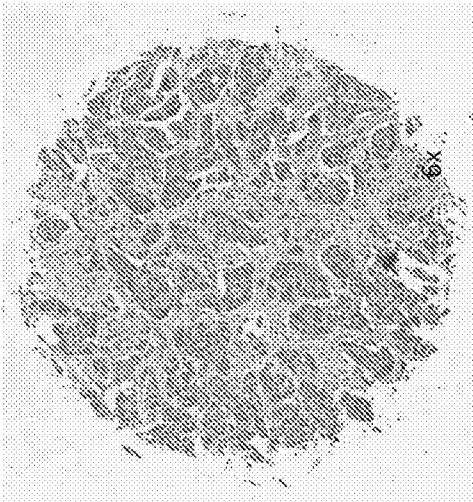
**Fig. 164A**

Position: A4  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



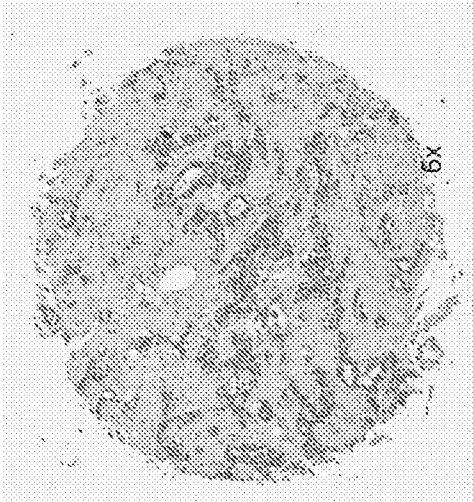
**Fig. 164B**

Position: A11  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0

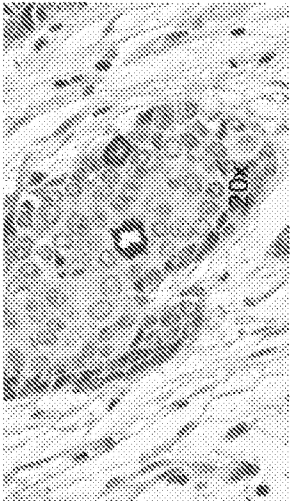


**Fig. 164C**

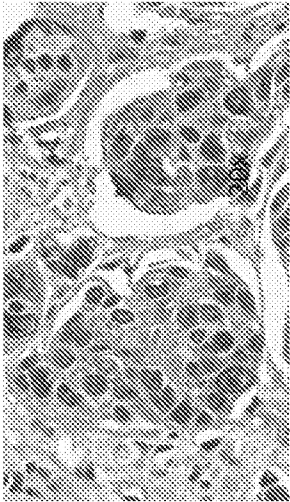
Position: G11  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



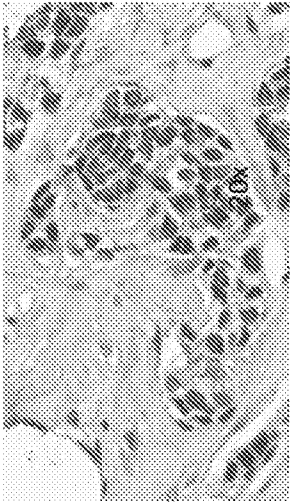
**Fig. 164D**



**Fig. 164E**



**Fig. 164F**



**Fig. 164A-164F**



**N+20 antibody 29H1 0.5 ug/mL**  
**Pancreatic cancer tissue array PA1003**

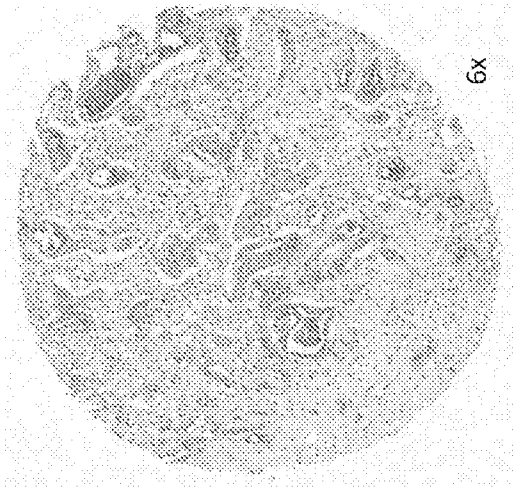
**Fig. 166A**

Position: A8  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



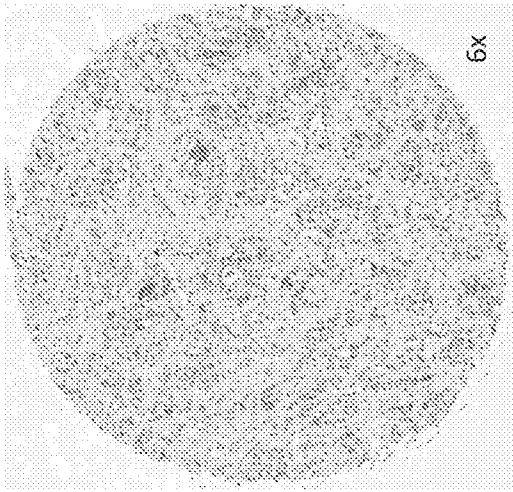
**Fig. 166B**

Position: B10  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0

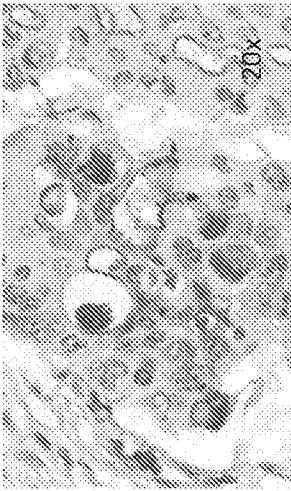


**Fig. 166C**

Position: D8  
Cell Type: Adenocarcinoma  
Tumor Grade: 3  
TNM: T3N0M0



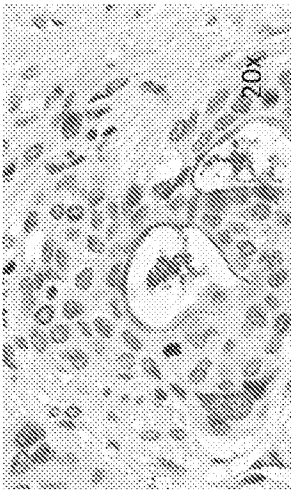
**Fig. 166D**



**Fig. 166E**



**Fig. 166F**



**Fig. 166A--166F**

N+20/C-27 antibody 31A1 IgG 0.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 167A

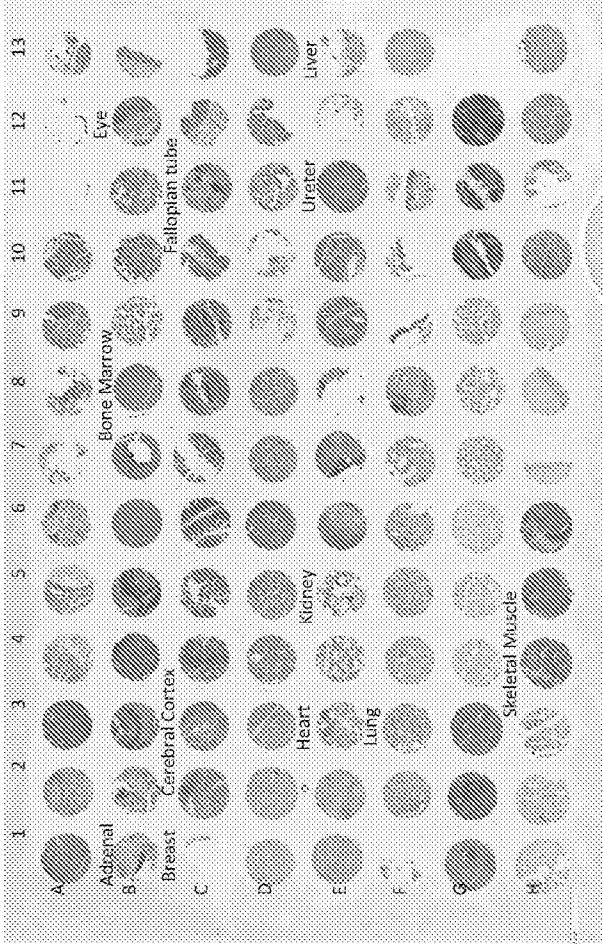


Fig. 167C

Position	Age	Sex	Organ/Anatomic Site	Pathology	Position	Age	Sex	Organ/Anatomic Site	Pathology
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	51	Adrenal gland	Normal	E3	M	72	Lung	Normal
A4	M	72	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	45	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	52	Bladder, urinary	Normal	E7	F	46	Ovary	Normal
A8	M	53	Bone, bone marrow	Normal	E8	F	46	Ovary	Normal
A9	F	44	Bone, bone marrow	Normal	E9	M	42	Parotids	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	26	Parotids	Normal
A11	M	55	Head and neck, salivary gland	Normal	E11	F	59	Parotids	Normal
A12	M	55	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	F	35	Eye	Normal	E13	M	72	Parathyroid	Adenoma
B1	F	35	Breast	Normal	F1	F	3	Pituitary gland	Normal
B2	F	35	Breast	Normal	F2	F	32	Pituitary gland	Normal
B3	M	85	Brain, cerebellum	Normal	F3	F	30	Pituitary gland	Normal
B4	M	85	Brain, cerebellum	Normal	F4	F	27	Pituitary gland	Normal
B5	M	85	Brain, cerebellum	Normal	F5	M	84	Prostate	Normal
B6	M	85	Brain, cerebral cortex	Normal	F6	M	85	Prostate	Normal
B7	M	85	Brain, cerebral cortex	Normal	F7	M	85	Prostate	Normal
B8	F	42	Brain, cerebral cortex	Normal	F8	M	85	Prostate	Normal
B9	F	42	Brain, cerebral cortex	Normal	F9	F	32	Skin	Normal
B10	F	32	Fallopian tube	Normal	F10	F	26	Skin	Normal
B11	F	24	Fallopian tube	Normal	F11	M	26	Skin	Normal
B12	M	45	Esophagus	Normal	F12	M	58	Spinal cord	Normal
B13	M	34	Esophagus	Normal	G1	M	27	Spleen	Normal
C1	M	45	Esophagus	Normal	G2	M	31	Spleen	Normal
C2	M	45	Esophagus	Normal	G3	M	31	Spleen	Normal
C3	M	45	Esophagus	Normal	G4	F	90	Skeletal muscle	Normal
C4	M	77	Stomach	Normal	G5	M	49	Skeletal muscle	Normal
C5	M	45	Intestine, small intestine	Normal	G6	M	70	Skeletal muscle	Normal
C6	F	75	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C7	F	75	Intestine, small intestine	Normal	G8	M	43	Testis	Normal
C8	M	2	Intestine, colon	Normal	G9	M	77	Testis	Normal
C9	M	46	Intestine, colon	Normal	G10	M	15	Thymus	Normal
C10	M	74	Intestine, colon	Normal	G11	M	26	Thymus	Normal
C11	F	77	Intestine, rectum	Normal	G12	F	9	Thymus	Normal
C12	F	77	Intestine, rectum	Normal	G13	F	26	Thyroid	Normal
C13	M	26	Intestine, rectum	Normal	H1	F	37	Thyroid	Normal
D1	M	35	Heart	Normal	H2	F	51	Thyroid	Normal
D2	M	35	Heart	Normal	H3	M	46	Tonsil	Normal
D3	F	44	Heart	Normal	H4	M	37	Tonsil	Normal
D4	F	52	Kidney, cortex	Normal	H5	M	13	Tonsil	Normal
D5	M	23	Kidney, cortex	Normal	H6	F	38	Uterus, cervix	Normal
D6	M	53	Kidney, medulla	Normal	H7	F	35	Uterus, cervix	Normal
D7	M	53	Kidney, medulla	Schwannoma	H8	F	44	Uterus, cervix	Normal
D8	M	53	Kidney, medulla	Schwannoma	H9	F	36	Uterus	Normal
D9	M	39	Peripheral nerve	Schwannoma	H10	F	36	Uterus	Normal
D10	F	28	Peripheral nerve	Schwannoma	H11	F	41	Uterus	Normal
D11	F	44	Ureter	Normal	H12	F	46	Uterus	Normal
D12	F	42	Ureter	Normal	H13	M	56	Skin	Malignant melanoma (tissue marker)
D13	F	57	Liver	Normal					

Fig. 167B

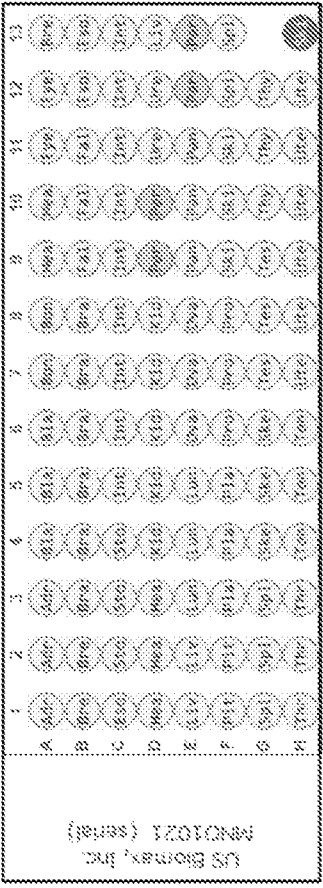


Figure 167A-167C



N+20/C-27 antibody 31A1 IgG 0.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 168A

Normal Adrenal Gland

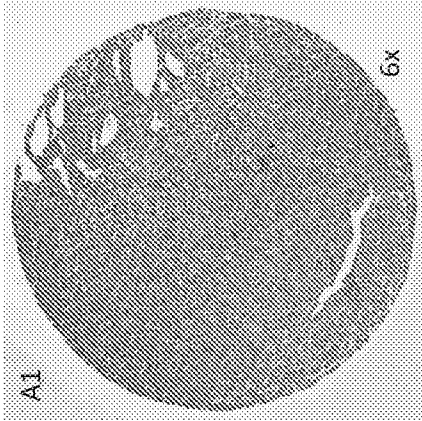


Fig. 168B

Normal Breast

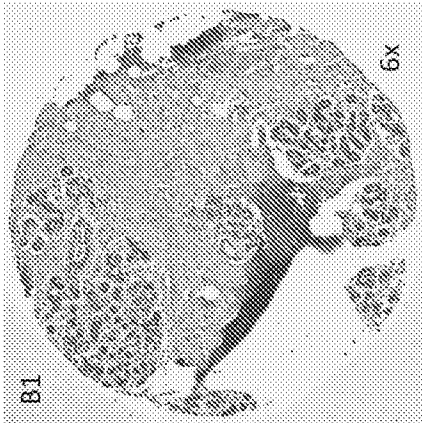


Fig. 168C

Normal Fallopian Tubes

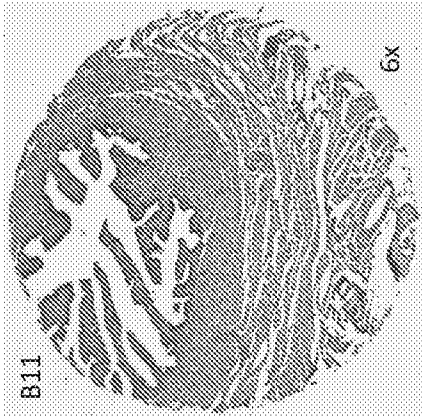


Fig. 168D

Normal Kidney

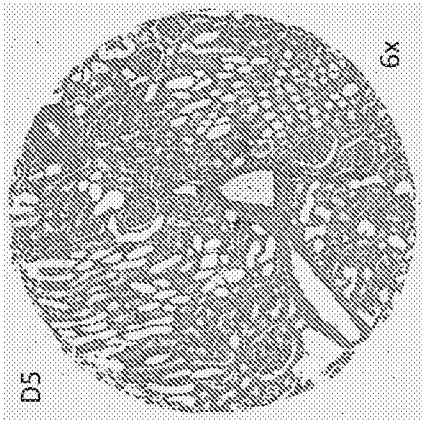


Fig. 168E

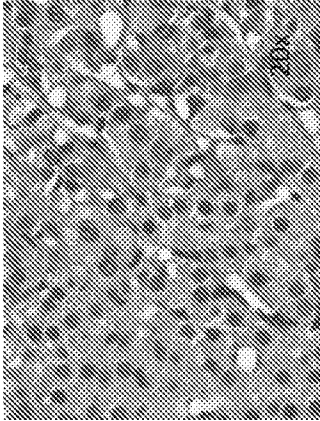


Fig. 168F



Fig. 168G



Fig. 168H

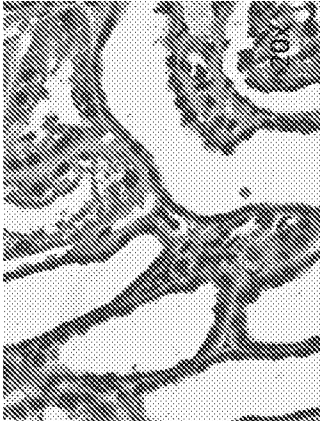


Figure 168A-168H

N+20/C-27 antibody 31A1 IgG 0.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 168I  
Normal Heart

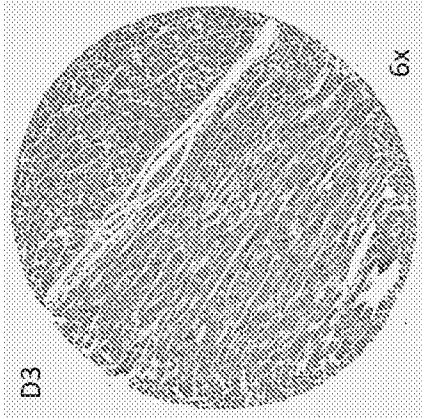


Fig. 168J  
Normal Liver

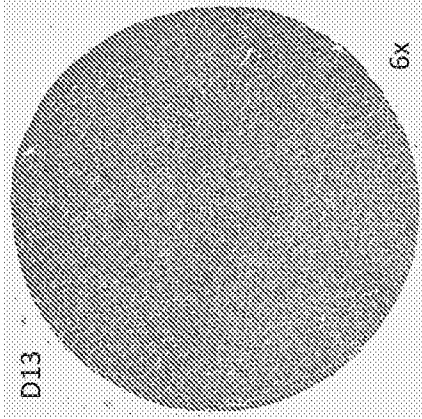


Fig. 168K  
Normal Lung

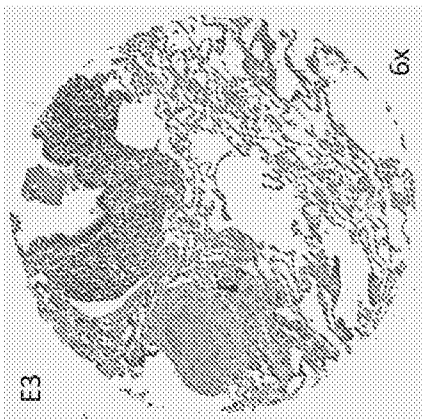


Fig. 168L  
Normal Ureter

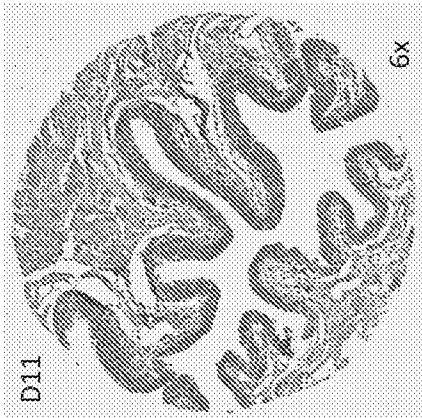


Fig. 168M



Fig. 168N

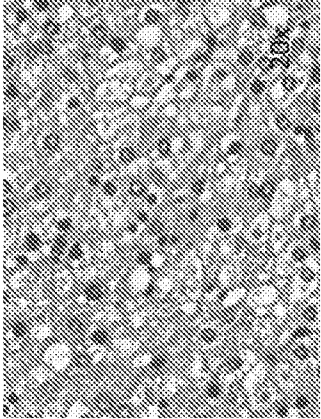


Fig. 168O



Fig. 168P

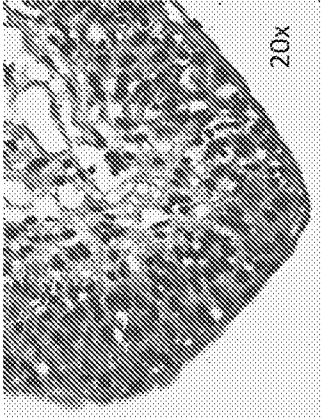


Figure 168I-168P



N+20/C-27 antibody 31A1 IgG 0.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 168Q  
Normal Eye

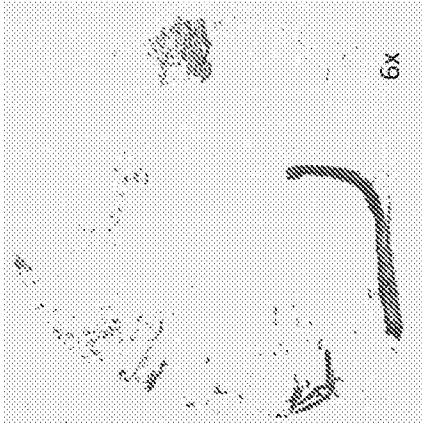


Fig. 168R  
Normal Cerebral Cortex

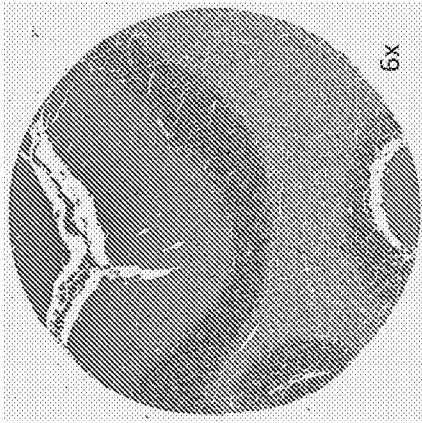


Fig. 168S  
Normal Bone Marrow

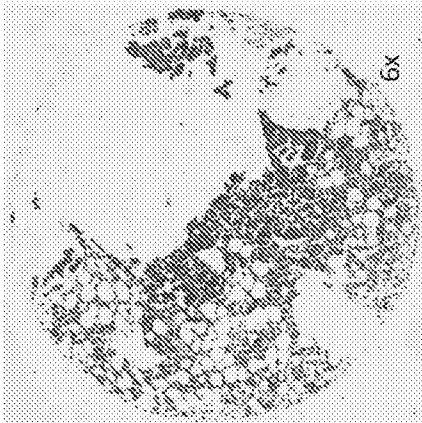


Fig. 168T  
Normal Skeletal Muscle

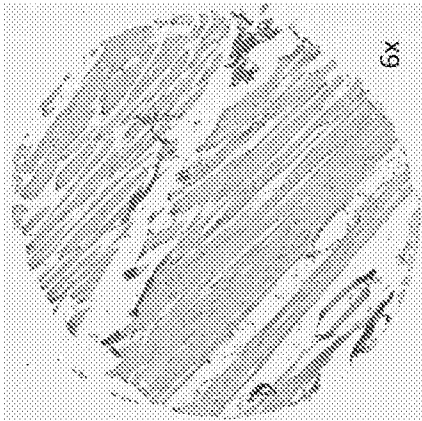


Fig. 168U

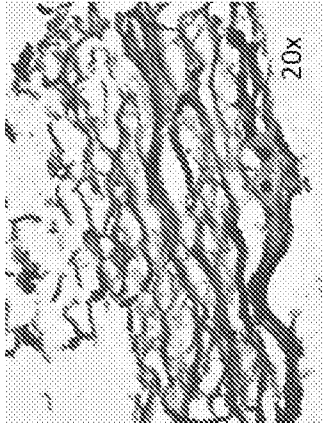


Fig. 168V

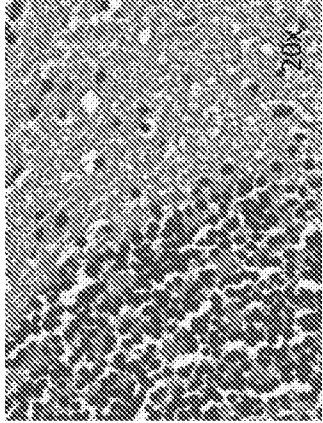


Fig. 168W

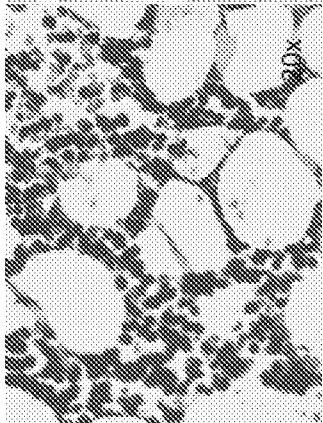


Fig. 168X

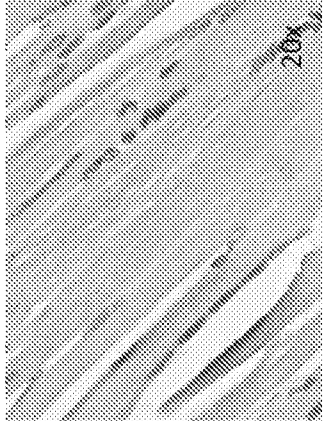


Figure 168Q-168X

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# N+20/C-27 antibody 31A1 IgG 0.5 ug/mL Breast cancer tissue array Br11141

Fig. 169A

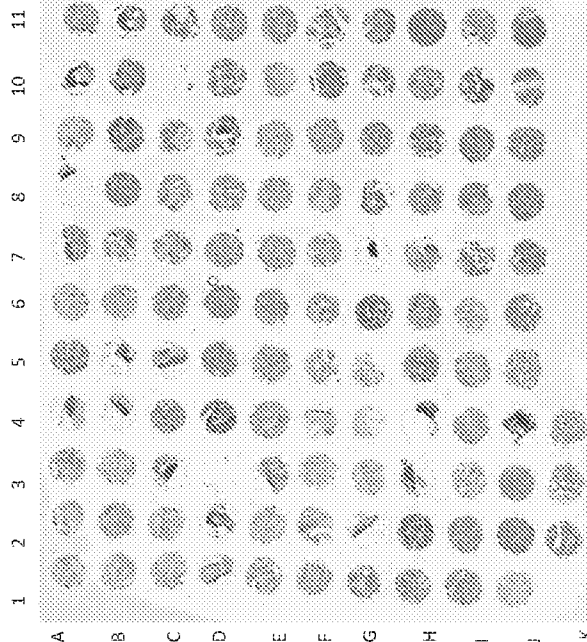


Fig. 169B

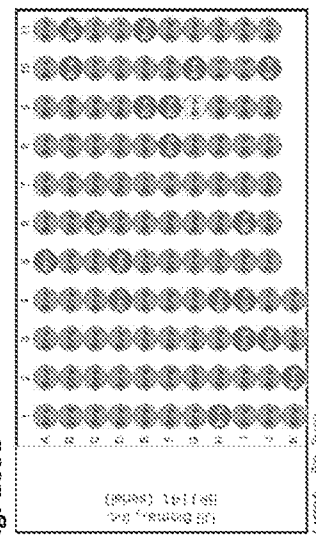


Figure 169A-169C

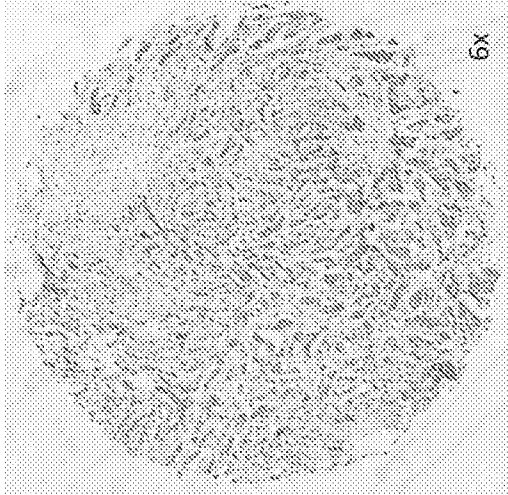
Fig. 169C

Protein	Age	Sex	Primary diagnosis	TNM	Stage	Type	Position	Site	Pathology	Notes	Stage	Type	
A1	45	F	Invasive ductal carcinoma	T2N0M0	2	IA	F1	52	F	Invasive ductal carcinoma	T2N0M0	2	IA
A2	38	F	Invasive ductal carcinoma	T2N0M0	2	IA	F2	44	F	Invasive ductal carcinoma	T2N0M0	2	IA
A3	42	F	Invasive ductal carcinoma	T2N0M0	2	IA	F3	53	F	Invasive ductal carcinoma	T2N0M0	2	IA
A4	40	F	Invasive ductal carcinoma	T2N0M0	2	IA	F4	54	F	Invasive ductal carcinoma	T2N1M0	2	IB
A5	70	F	Invasive ductal carcinoma	T2N1M0	2	IA	F5	53	F	Invasive ductal carcinoma	T2N1M0	2	IB
A6	48	F	Invasive ductal carcinoma	T2N0M0	2	IB	F6	59	F	Invasive ductal carcinoma	T2N1M0	2	IB
A7	42	F	Invasive ductal carcinoma	T2N1M0	2	IB	F7	43	F	Invasive ductal carcinoma	T2N1M0	2	IB
A8	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	F8	45	F	Invasive ductal carcinoma	T2N1M0	2	IB
A9	58	F	Invasive ductal carcinoma	T2N1M0	2	IB	F9	52	F	Invasive ductal carcinoma	T2N1M0	2	IB
A10	52	F	Invasive ductal carcinoma	T2N0M0	2	IB	F10	42	F	Invasive ductal carcinoma	T2N1M0	2	IB
A11	50	F	Invasive ductal carcinoma	T2N0M0	2	IA	F11	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
B1	43	F	Invasive ductal carcinoma	T2N0M0	2	IB	G1	45	F	Invasive ductal carcinoma	T2N0M0	2	IB
B2	37	F	Invasive ductal carcinoma	T2N1M0	2	IB	G2	42	F	Invasive ductal carcinoma	T2N0M0	2	IB
B3	57	F	Invasive ductal carcinoma	T2N0M0	2	IA	G3	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
B4	77	F	Invasive ductal carcinoma	T2N0M0	2	IA	G4	45	F	Invasive ductal carcinoma	T2N0M0	2	IB
B5	58	F	Invasive ductal carcinoma	T2N0M0	2	IA	G5	45	F	Invasive ductal carcinoma	T2N0M0	2	IA
B6	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	G6	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
B7	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	G7	53	F	Invasive ductal carcinoma	T2N0M0	2	IA
B8	50	F	Invasive ductal carcinoma	T2N0M0	2	IA	G8	42	F	Invasive ductal carcinoma	T2N1M0	2	IB
B9	50	F	Invasive ductal carcinoma	T2N0M0	2	IA	G9	38	F	Invasive ductal carcinoma	T2N0M0	2	IB
B10	40	F	Invasive ductal carcinoma	T2N1M0	2	IB	G10	46	F	Invasive ductal carcinoma	T2N1M0	2	IB
B11	47	F	Invasive ductal carcinoma	T2N1M0	2	IB	G11	47	F	Invasive ductal carcinoma	T2N1M0	2	IB
C1	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	G12	41	F	Invasive ductal carcinoma	T2N1M0	2	IB
C2	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	H1	45	F	Invasive ductal carcinoma	T2N1M0	2	IB
C3	42	F	Invasive ductal carcinoma	T2N0M0	2	IA	H2	53	F	Invasive ductal carcinoma	T2N0M0	2	IB
C4	41	F	Invasive ductal carcinoma	T2N0M0	2	IA	H3	52	F	Invasive ductal carcinoma	T2N0M0	2	IA
C5	55	F	Invasive ductal carcinoma	T2N0M0	2	IA	H4	50	F	Invasive ductal carcinoma	T2N1M0	2	IB
C6	41	F	Invasive ductal carcinoma	T2N1M0	2	IB	H5	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
C7	41	F	Invasive ductal carcinoma	T2N0M0	2	IB	H6	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
C8	40	F	Invasive ductal carcinoma	T2N0M0	2	IB	H7	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
C9	40	F	Invasive ductal carcinoma	T2N0M0	2	IB	H8	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
C10	42	F	Invasive ductal carcinoma	T2N1M0	2	IB	H9	41	F	Invasive ductal carcinoma	T2N0M0	2	IA
C11	57	F	Invasive ductal carcinoma	T2N0M0	2	IA	H10	51	F	Invasive ductal carcinoma	T2N0M0	2	IA
C12	49	F	Invasive ductal carcinoma	T2N0M0	2	IA	H11	42	F	Invasive ductal carcinoma	T2N0M0	2	IB
D1	49	F	Invasive ductal carcinoma	T2N0M0	2	IA	I1	45	F	Invasive ductal carcinoma	T2N1M0	2	IB
D2	48	F	Invasive ductal carcinoma	T2N0M0	2	IB	I2	66	F	Invasive ductal carcinoma	T2N0M0	2	IB
D3	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	I3	75	F	Invasive ductal carcinoma	T2N1M0	2	IB
D4	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	I4	58	F	Invasive ductal carcinoma	T2N1M0	2	IB
D5	52	F	Invasive ductal carcinoma	T2N1M0	2	IB	I5	54	F	Invasive ductal carcinoma	T2N1M0	2	IB
D6	52	F	Invasive ductal carcinoma	T2N0M0	2	IA	I6	54	F	Invasive ductal carcinoma	T2N0M0	2	IB
D7	52	F	Invasive ductal carcinoma	T2N0M0	2	IA	I7	52	F	Invasive ductal carcinoma	T2N0M0	2	IB
D8	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	I8	52	F	Invasive ductal carcinoma	T2N0M0	2	IB
D9	43	F	Invasive ductal carcinoma	T2N1M0	2	IB	I9	59	F	Invasive ductal carcinoma	T2N0M0	2	IA
D10	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	I10	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
D11	50	F	Invasive ductal carcinoma	T2N0M0	2	IA	I11	57	F	Invasive ductal carcinoma	T2N0M0	2	IA
E1	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J1	73	F	Invasive ductal carcinoma	T2N0M0	2	IA
E2	57	F	Invasive ductal carcinoma	T2N0M0	2	IA	J2	66	F	Invasive ductal carcinoma	T2N0M0	2	IA
E3	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	J3	45	F	Invasive ductal carcinoma	T2N0M0	2	IA
E4	44	F	Invasive ductal carcinoma	T2N0M0	2	IB	J4	44	F	Invasive ductal carcinoma	T2N0M0	2	IB
E5	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J5	48	F	Invasive ductal carcinoma	T2N0M0	2	IB
E6	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J6	52	F	Invasive ductal carcinoma	T2N0M0	2	IB
E7	58	F	Invasive ductal carcinoma	T2N0M0	2	IA	J7	56	F	Invasive ductal carcinoma	T2N0M0	2	IA
E8	58	F	Invasive ductal carcinoma	T2N0M0	2	IA	J8	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
E9	58	F	Invasive ductal carcinoma	T2N1M0	2	IB	J9	51	F	Invasive ductal carcinoma	T2N1M0	2	IB
E10	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	J10	53	F	Invasive ductal carcinoma	T2N1M0	2	IB
E11	52	F	Invasive ductal carcinoma	T2N1M0	2	IB	J11	57	F	Invasive ductal carcinoma	T2N0M0	2	IB
						K1	45	F	Invasive ductal carcinoma	T2N0M0	2	IB	
						K2	42	F	Invasive ductal carcinoma	T2N1M0	2	IB	
						K3	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	
						K4	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	

**N+20/C-27 antibody 31A1 IgG 15 ug/mL**  
**Breast cancer tissue array Br1141**

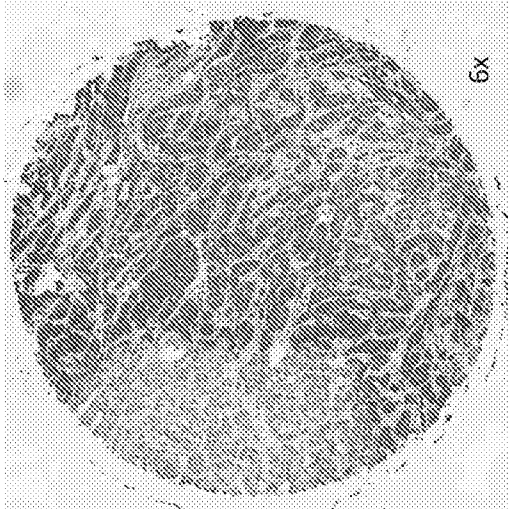
**Fig. 170A**

Position: A4  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



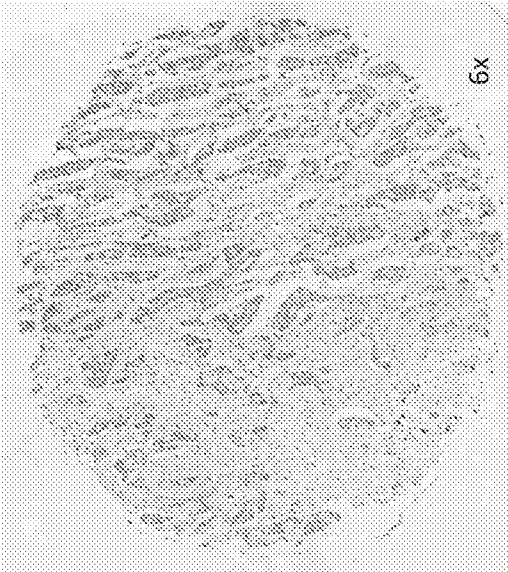
**Fig. 170B**

Position: A11  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0

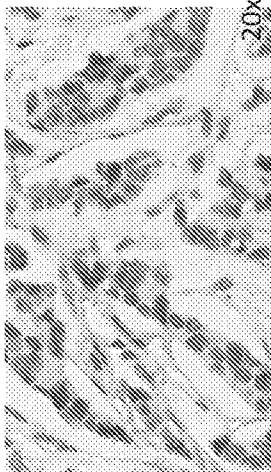


**Fig. 170C**

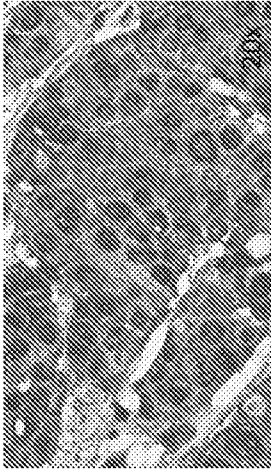
Position: G11  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



**Fig. 170D**



**Fig. 170E**



**Fig. 170F**



**Fig. 170A-170F**

N+20 antibody 31A1 0.1 - 10 ug/ml  
Pancreatic cancer tissue array PA1003

Fig. 171A

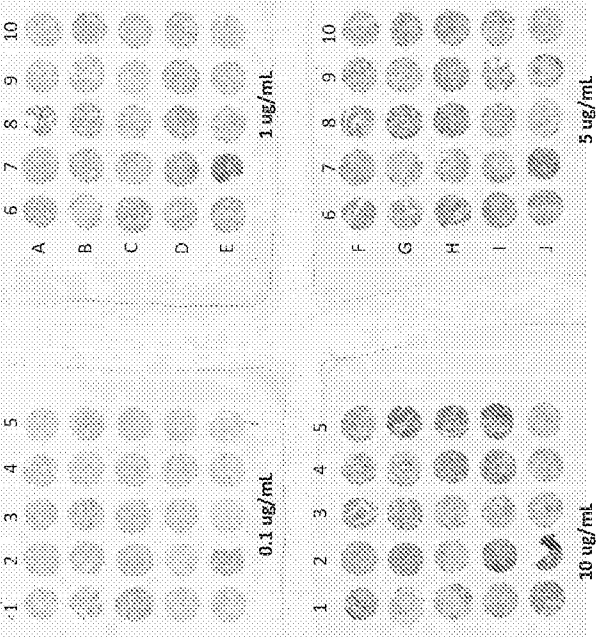


Fig. 171B



Fig. 171A-171C

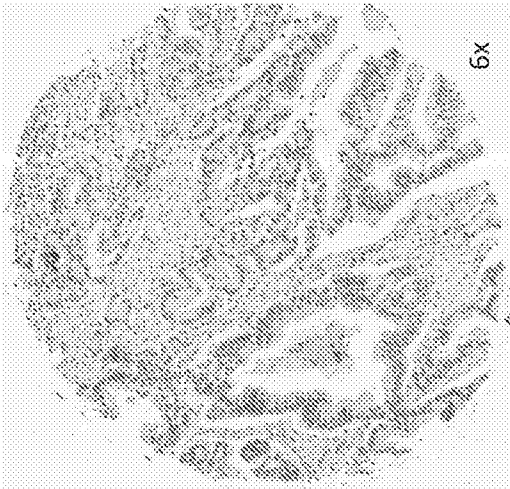
Fig. 171C

A1	38	F	Adenocarcinoma	TSN000	2	II	71	35	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A2	41	M	Adenocarcinoma	TSN000	2	II	53	44	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A3	59	M	Adenocarcinoma	TSN000	2	II	53	35	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A4	65	M	Adenocarcinoma	TSN000	2	II	54	65	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A5	53	M	Adenocarcinoma	TSN000	2	II	55	53	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A6	39	F	Adenocarcinoma	TSN000	2	II	55	35	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A7	44	M	Adenocarcinoma	TSN000	2	II	77	44	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A8	59	M	Adenocarcinoma	TSN000	2	II	78	59	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A9	65	M	Adenocarcinoma	TSN000	2	II	79	65	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
A10	53	M	Adenocarcinoma	TSN000	2	II	79	53	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B1	42	M	Adenocarcinoma	TSN001	2	IV	710	42	M	Adenocarcinoma	TSN001	2	IV	malignant	TSN001	2	IV	malignant
B2	52	M	Adenocarcinoma	TSN000	2	II	710	52	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B3	57	M	Adenocarcinoma	TSN000	2	II	710	57	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B4	34	M	Adenocarcinoma	TSN000	2	II	710	34	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B5	71	F	Adenocarcinoma	TSN000	2	II	710	71	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B6	42	M	Adenocarcinoma	TSN001	2	IV	710	42	M	Adenocarcinoma	TSN001	2	IV	malignant	TSN001	2	IV	malignant
B7	52	M	Adenocarcinoma	TSN000	2	II	710	52	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B8	57	M	Adenocarcinoma	TSN000	2	II	710	57	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B9	34	M	Adenocarcinoma	TSN000	2	II	710	34	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
B10	72	F	Adenocarcinoma	TSN000	2	II	710	72	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C1	47	M	Adenocarcinoma	TSN000	2	II	710	47	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C2	64	F	Adenocarcinoma	TSN000	2	II	710	64	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C3	51	M	Adenocarcinoma	TSN000	2	II	710	51	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C4	68	F	Adenocarcinoma	TSN000	2	II	710	68	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C5	41	M	Adenocarcinoma	TSN000	2	II	710	41	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C6	47	M	Adenocarcinoma	TSN000	2	II	710	47	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C7	64	F	Adenocarcinoma	TSN000	2	II	710	64	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C8	51	M	Adenocarcinoma	TSN000	2	II	710	51	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C9	68	F	Adenocarcinoma	TSN000	2	II	710	68	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
C10	41	M	Adenocarcinoma	TSN000	2	II	710	41	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D1	58	F	Adenocarcinoma	TSN000	2	II	710	58	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D2	41	F	Adenocarcinoma	TSN000	2	II	710	41	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D3	77	M	Adenocarcinoma	TSN000	2	II	710	77	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D4	59	M	Adenocarcinoma	TSN000	2	II	710	59	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D5	56	F	Adenocarcinoma	TSN000	2	II	710	56	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D6	58	F	Adenocarcinoma	TSN000	2	II	710	58	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D7	41	F	Adenocarcinoma	TSN000	2	II	710	41	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D8	72	F	Adenocarcinoma	TSN000	2	II	710	72	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D9	59	M	Adenocarcinoma	TSN000	2	II	710	59	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
D10	59	M	Adenocarcinoma	TSN000	2	II	710	59	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E1	53	F	Adenocarcinoma	TSN000	2	II	710	53	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E2	60	M	Adenocarcinoma	TSN000	2	II	710	60	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E3	56	M	Adenocarcinoma	TSN000	2	II	710	56	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E4	40	F	Adenocarcinoma	TSN000	2	II	710	40	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E5	46	F	Adenocarcinoma	TSN000	2	II	710	46	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E6	55	F	Adenocarcinoma	TSN000	2	II	710	55	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E7	60	M	Adenocarcinoma	TSN000	2	II	710	60	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E8	56	M	Adenocarcinoma	TSN000	2	II	710	56	M	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E9	60	F	Adenocarcinoma	TSN000	2	II	710	60	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant
E10	66	F	Adenocarcinoma	TSN000	2	II	710	66	F	Adenocarcinoma	TSN000	2	II	malignant	TSN000	2	II	malignant

**N+20/C-27-antibody 31A1 1 ug/mL**  
**Pancreatic cancer tissue array PA1003**

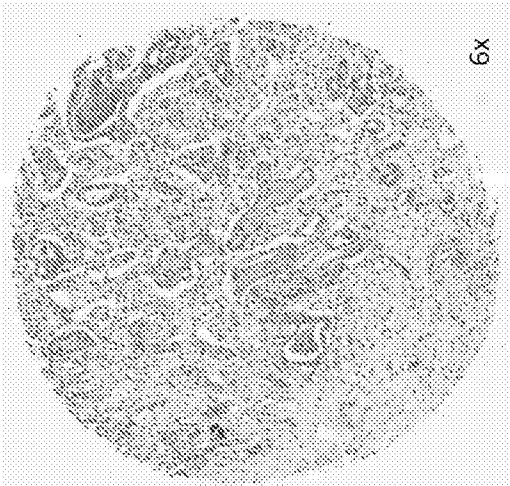
**Fig. 172A**

Position: B7  
Cell Type: Adenocarcinoma  
Tumor Grade: 1  
TNM: T3N0M0



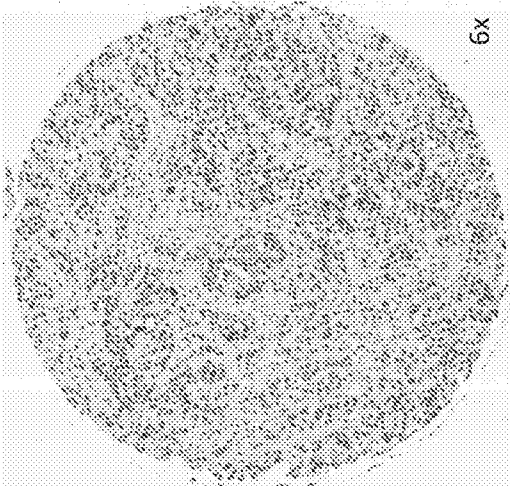
**Fig. 172B**

Position: B10  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0

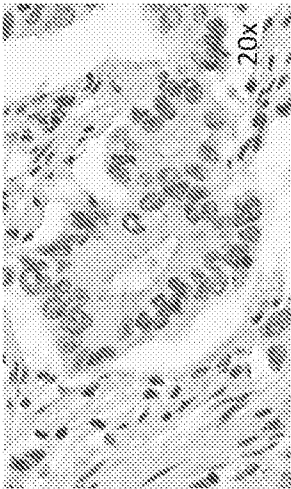


**Fig. 172C**

Position: D8  
Cell Type: Adenocarcinoma  
Tumor Grade: 3  
TNM: T3N0M0



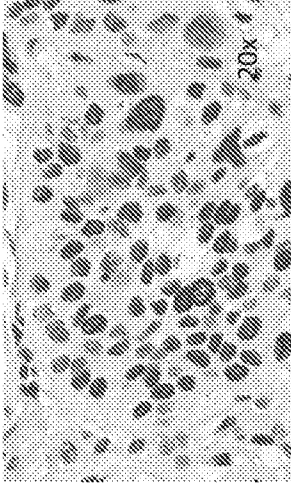
**Fig. 172D**



**Fig. 172E**



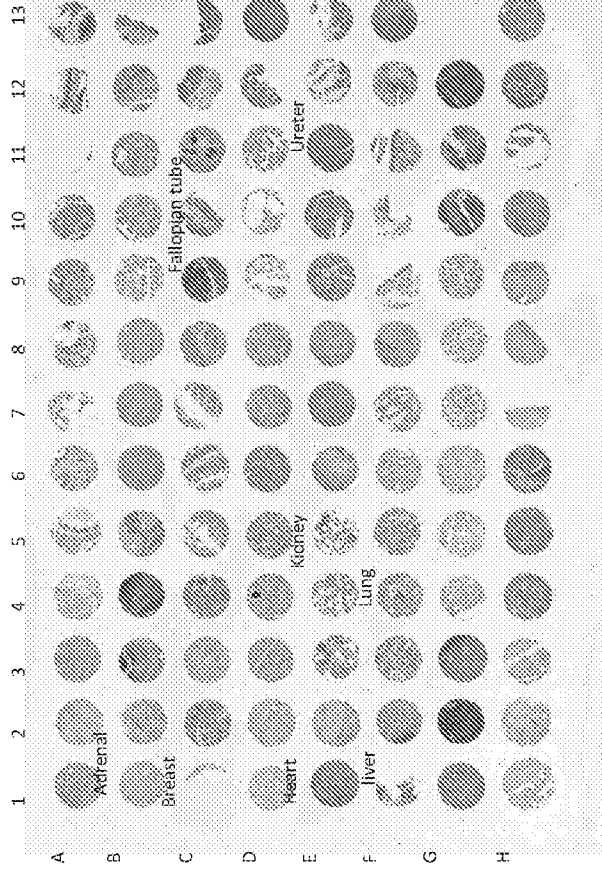
**Fig. 172F**



**Fig. 172A-172F**

**N+20/C-27 antibody 32C1 IgG 0.25 ug/mL**  
**FDA Normal tissue array MNO1021**

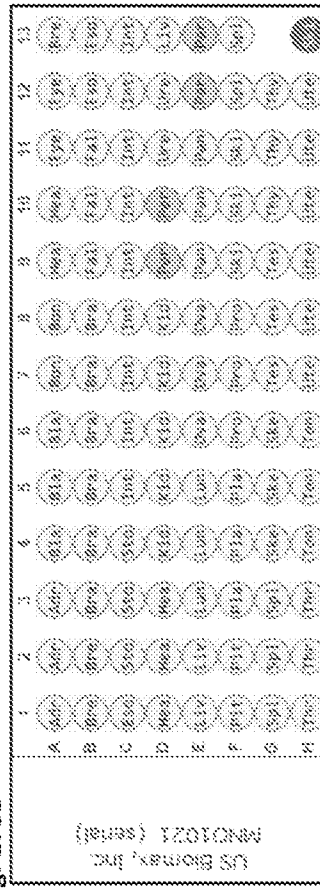
**Fig. 173A**



**Fig. 173C**

Position	Age	Sex	Organ/Anatomic Site	Pathology	Position	Age	Sex	Organ/Anatomic Site	Pathology
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	51	Adrenal gland	Normal	E3	M	72	Liver	Normal
A4	M	72	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	45	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	52	Bladder, urinary	Normal	E7	F	46	Ovary	Normal
A8	M	53	Bone, bone marrow	Normal	E8	F	46	Ovary	Normal
A9	F	44	Bone, bone marrow	Normal	E9	M	42	Parotid	Normal
A10	F	18	Head and neck, salivary gland	Normal	E10	F	26	Parotid	Normal
A11	M	55	Head and neck, salivary gland	Normal	E11	F	59	Parotid	Normal
A12	M	50	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	F	35	Breast	Normal	E13	F	72	Parathyroid	Adenoma
B1	F	38	Breast	Normal	F1	F	3	Mammary gland	Normal
B2	F	38	Breast	Normal	F2	F	32	Mammary gland	Normal
B3	M	85	Brain, cerebellum	Normal	F3	F	30	Placenta	Normal
B4	M	85	Brain, cerebellum	Normal	F4	F	27	Placenta	Normal
B5	M	85	Brain, cerebral cortex	Normal	F5	M	84	Prostate	Normal
B6	M	85	Brain, cerebral cortex	Normal	F6	M	85	Prostate	Normal
B7	F	42	Brain, cerebral cortex	Normal	F7	M	85	Prostate	Normal
B8	F	42	Brain, cerebral cortex	Normal	F8	M	85	Prostate	Normal
B9	F	32	Fallopian tube	Normal	F9	F	32	Skin	Normal
B10	F	32	Fallopian tube	Normal	F10	F	32	Skin	Normal
B11	F	24	Fallopian tube	Normal	F11	M	26	Skin	Normal
B12	M	34	Esophagus	Normal	F12	M	57	Spinal cord	Normal
B13	M	34	Esophagus	Normal	F13	M	57	Spinal cord	Normal
C1	M	34	Esophagus	Normal	G1	M	27	Spleen	Normal
C2	M	45	Esophagus	Normal	G2	M	31	Spleen	Normal
C3	M	45	Esophagus	Normal	G3	M	31	Spleen	Normal
C4	M	77	Stomach	Normal	G4	F	90	Skeletal muscle	Normal
C5	M	77	Stomach	Normal	G5	M	49	Skeletal muscle	Normal
C6	M	45	Intestine, small intestine	Normal	G6	M	70	Skeletal muscle	Normal
C7	M	75	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C8	M	75	Intestine, small intestine	Normal	G8	M	43	Testis	Normal
C9	M	75	Intestine, small intestine	Normal	G9	M	43	Testis	Normal
C10	M	75	Intestine, small intestine	Normal	G10	M	15	Thymus	Normal
C11	M	75	Intestine, small intestine	Normal	G11	M	26	Thymus	Normal
C12	M	75	Intestine, small intestine	Normal	G12	F	9	Thymus	Normal
C13	M	75	Intestine, small intestine	Normal	G13	F	9	Thymus	Normal
D1	M	28	Heart	Normal	H1	F	26	Thyroid	Normal
D2	M	28	Heart	Normal	H2	F	26	Thyroid	Normal
D3	M	28	Heart	Normal	H3	F	26	Thyroid	Normal
D4	M	28	Heart	Normal	H4	F	26	Thyroid	Normal
D5	M	23	Kidney, cortex	Normal	H5	M	46	Tonsil	Normal
D6	M	23	Kidney, cortex	Normal	H6	M	46	Tonsil	Normal
D7	M	23	Kidney, cortex	Normal	H7	M	46	Tonsil	Normal
D8	M	23	Kidney, cortex	Normal	H8	M	46	Tonsil	Normal
D9	M	23	Kidney, cortex	Normal	H9	M	46	Tonsil	Normal
D10	M	23	Kidney, cortex	Normal	H10	M	46	Tonsil	Normal
D11	M	23	Kidney, cortex	Normal	H11	M	46	Tonsil	Normal
D12	M	23	Kidney, cortex	Normal	H12	M	46	Tonsil	Normal
D13	M	23	Kidney, cortex	Normal	H13	M	46	Tonsil	Normal

**Fig. 173B**



**Figure 173A-173C**



N+20/C-27 antibody 32C1 IgG 0.25 ug/mL  
FDA Normal tissue array MNO1021

Fig. 174A  
Normal Adrenal Gland

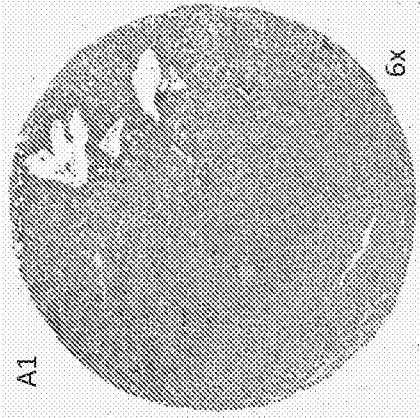


Fig. 174B  
Normal Breast

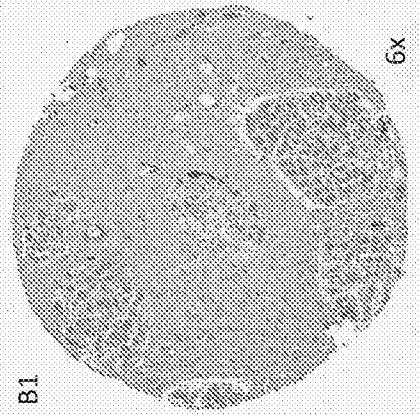


Fig. 174C  
Normal Fallopian Tubes

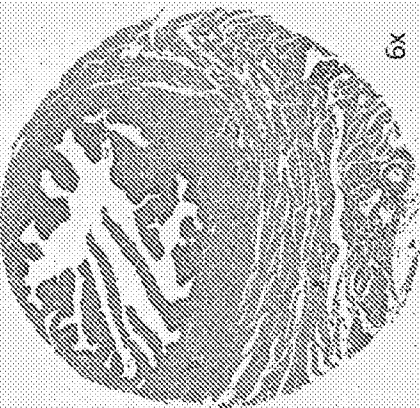


Fig. 174D  
Normal Kidney

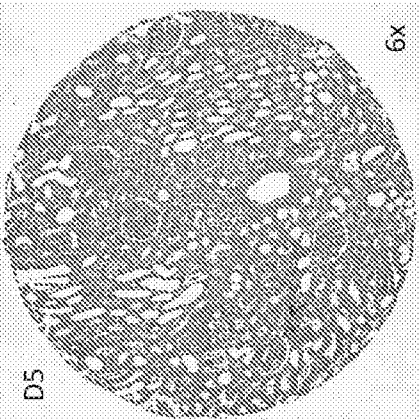


Fig. 174E

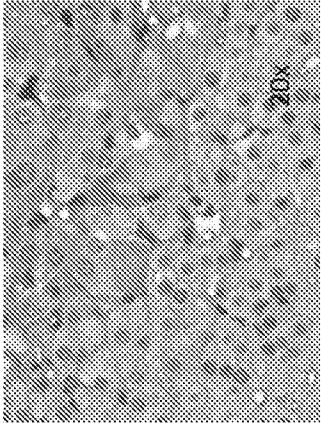


Fig. 174F



Fig. 174G

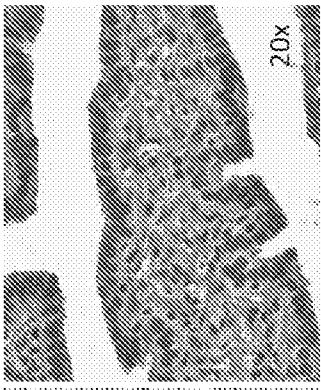


Fig. 174H

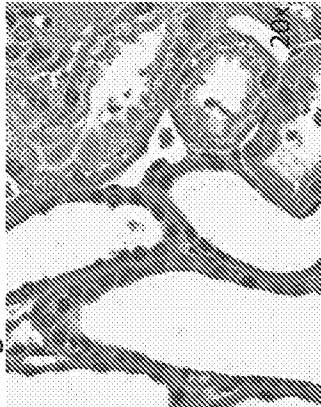


Figure 174A-174H

N+20/C-27 antibody 32C1 IgG 0.25 ug/mL  
FDA Normal tissue array MNO1021

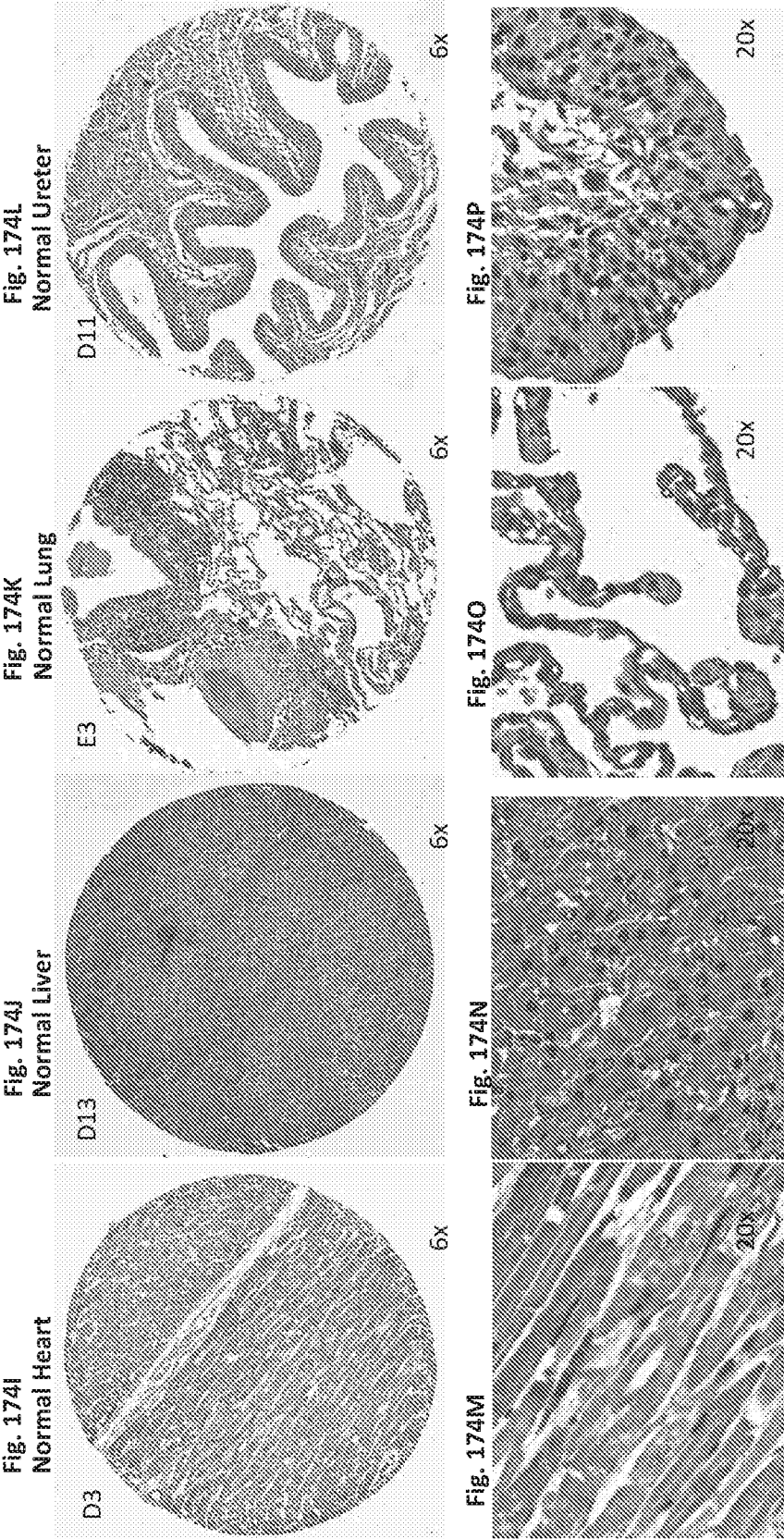


Figure 174I-174P



N+20/C-27 antibody 32C1 IgG 0.25 ug/mL  
FDA Normal tissue array MNO1021

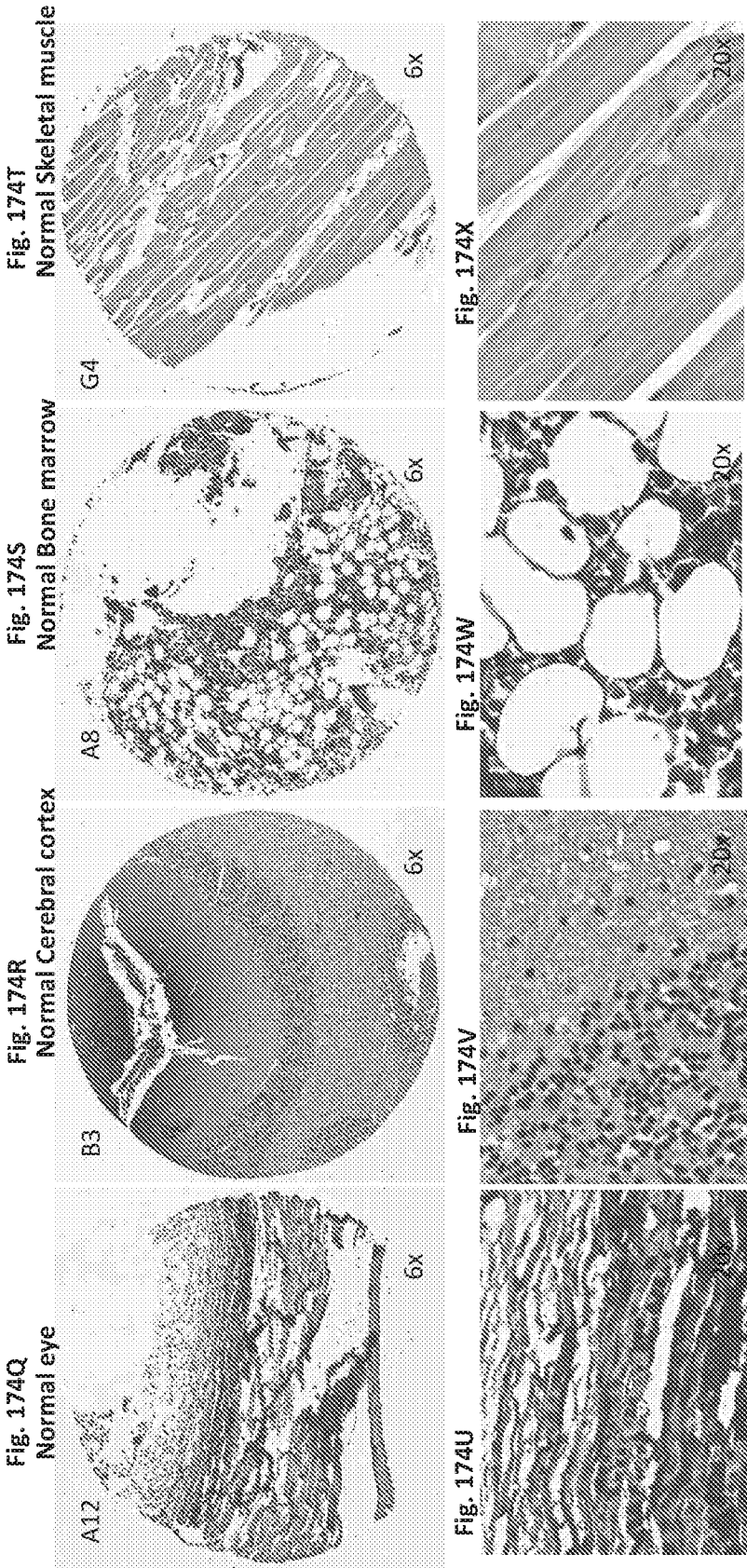


Figure 174Q-174X

N+20/C-27 antibody 32C1 IgG 0.25 ug/mL  
Breast cancer tissue array Br1141

Fig. 175A

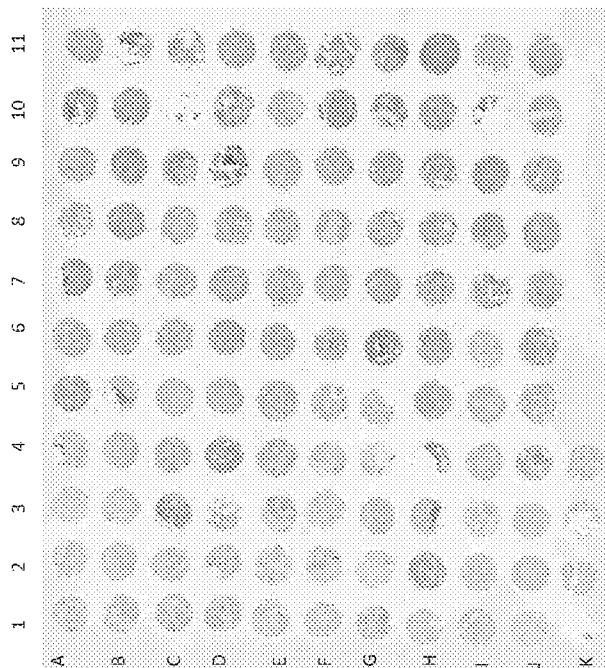


Fig. 175C

Position	Age	Sex	Primary diagnosis	TNM	Stages	Type	Position	Age	Sex	Primary diagnosis	TNM	Stages	Type
A1	45	F	Invasive ductal carcinoma	T2N0M0	2	IA	F1	52	F	Invasive ductal carcinoma	T2N0M0	2	IA
A2	38	F	Invasive ductal carcinoma	T2N0M0	2	IA	F2	44	F	Invasive ductal carcinoma	T2N0M0	2	IA
A3	42	F	Invasive ductal carcinoma	T2N0M0	2	IA	F3	53	F	Invasive ductal carcinoma	T2N0M0	2	IA
A4	40	F	Invasive ductal carcinoma	T2N0M0	2	IA	F4	50	F	Invasive ductal carcinoma	T2N0M0	2	IA
A5	70	F	Invasive ductal carcinoma	T2N1M0	2	IA	F5	53	F	Invasive ductal carcinoma	T2N1M0	2	IA
A6	48	F	Invasive ductal carcinoma	T2N0M0	2	IB	F6	59	F	Invasive ductal carcinoma	T2N1M0	2	IB
A7	42	F	Invasive ductal carcinoma	T2N1M0	2	IB	F7	43	F	Invasive ductal carcinoma	T2N0M0	2	IA
A8	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	F8	45	F	Invasive ductal carcinoma	T2N1M0	2	IA
A9	58	F	Invasive ductal carcinoma	T2N1M0	2	IB	F9	52	F	Invasive ductal carcinoma	T2N1M0	2	IB
A10	52	F	Invasive ductal carcinoma	T2N0M0	2	IB	F10	42	F	Invasive ductal carcinoma	T2N0M0	2	IB
A11	50	F	Invasive ductal carcinoma	T2N0M0	2	IB	F11	42	F	Invasive ductal carcinoma	T2N0M0	2	IB
B1	43	F	Invasive ductal carcinoma	T2N0M0	2	IA	G1	45	F	Invasive ductal carcinoma	T2N0M0	2	IA
B2	53	F	Invasive ductal carcinoma	T2N1M0	2	IB	G3	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
B3	57	F	Invasive ductal carcinoma	T2N1M0	2	IB	G4	45	F	Invasive ductal carcinoma	T2N0M0	2	IB
B4	77	F	Invasive ductal carcinoma	T2N0M0	2	IA	G4	45	F	Invasive ductal carcinoma	T2N0M0	2	IA
B5	58	F	Invasive ductal carcinoma	T2N0M0	2	IA	G5	45	F	Invasive ductal carcinoma	T2N0M0	2	IA
B6	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	G6	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
B7	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	G7	53	F	Invasive ductal carcinoma	T2N0M0	2	IA
B8	40	F	Invasive ductal carcinoma	T2N0M0	2	IA	G8	42	F	Invasive ductal carcinoma	T2N1M0	2	IB
B9	50	F	Invasive ductal carcinoma	T2N0M0	2	IA	G9	38	F	Invasive ductal carcinoma	T2N0M0	2	IB
B10	47	F	Invasive ductal carcinoma	T2N1M0	2	IB	G10	46	F	Invasive ductal carcinoma	T2N1M0	2	IB
B11	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	G11	45	F	Invasive ductal carcinoma	T2N1M0	2	IB
C1	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	G12	53	F	Invasive ductal carcinoma	T2N0M0	2	IB
C2	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	H3	52	F	Invasive ductal carcinoma	T2N0M0	2	IB
C3	42	F	Invasive ductal carcinoma	T2N0M0	2	IA	H4	50	F	Invasive ductal carcinoma	T2N1M0	2	IB
C4	41	F	Invasive ductal carcinoma	T2N0M0	2	IA	H5	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
C5	58	F	Invasive ductal carcinoma	T2N0M0	2	IA	H6	42	F	Invasive ductal carcinoma	T2N0M0	2	IA
C6	41	F	Invasive ductal carcinoma	T2N1M0	2	IB	H7	47	F	Invasive ductal carcinoma	T2N0M0	2	IA
C7	41	F	Invasive ductal carcinoma	T2N0M0	2	IB	H8	47	F	Invasive ductal carcinoma	T2N0M0	2	IB
C8	40	F	Invasive ductal carcinoma	T2N0M0	2	IB	H9	46	F	Invasive ductal carcinoma	T2N0M0	2	IB
C9	42	F	Invasive ductal carcinoma	T2N1M0	2	IB	H10	41	F	Invasive ductal carcinoma	T2N0M0	2	IB
C10	42	F	Invasive ductal carcinoma	T2N1M0	2	IB	H11	42	F	Invasive ductal carcinoma	T2N0M0	2	IB
C11	57	F	Invasive ductal carcinoma	T2N0M0	2	IA	I1	45	F	Invasive ductal carcinoma	T2N0M0	2	IB
C12	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	I2	66	F	Invasive ductal carcinoma	T2N0M0	2	IB
C13	42	F	Invasive ductal carcinoma	T2N0M0	2	IA	I3	75	F	Invasive ductal carcinoma	T2N1M0	2	IB
C14	41	F	Invasive ductal carcinoma	T2N0M0	2	IA	I4	58	F	Invasive ductal carcinoma	T2N1M0	2	IB
C15	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	I5	54	F	Invasive ductal carcinoma	T2N0M0	2	IA
C16	52	F	Invasive ductal carcinoma	T2N1M0	2	IB	I6	52	F	Invasive ductal carcinoma	T2N0M0	2	IB
C17	52	F	Invasive ductal carcinoma	T2N0M0	2	IA	I7	52	F	Invasive ductal carcinoma	T2N0M0	2	IB
C18	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	I8	52	F	Invasive ductal carcinoma	T2N0M0	2	IB
C19	43	F	Invasive ductal carcinoma	T2N1M0	2	IB	I9	59	F	Invasive ductal carcinoma	T2N0M0	2	IB
C20	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	I10	47	F	Invasive ductal carcinoma	T2N0M0	2	IB
D1	40	F	Invasive ductal carcinoma	T2N0M0	2	IA	I11	57	F	Invasive ductal carcinoma	T2N0M0	2	IB
D2	48	F	Invasive ductal carcinoma	T2N0M0	2	IB	J1	73	F	Invasive ductal carcinoma	T2N0M0	2	IB
D3	48	F	Invasive ductal carcinoma	T2N0M0	2	IB	J2	66	F	Invasive ductal carcinoma	T2N0M0	2	IB
D4	48	F	Invasive ductal carcinoma	T2N1M0	2	IB	J3	45	F	Invasive ductal carcinoma	T2N0M0	2	IB
D5	52	F	Invasive ductal carcinoma	T2N1M0	2	IB	J4	44	F	Invasive ductal carcinoma	T2N0M0	2	IB
D6	52	F	Invasive ductal carcinoma	T2N0M0	2	IB	J5	48	F	Invasive ductal carcinoma	T2N0M0	2	IB
D7	52	F	Invasive ductal carcinoma	T2N0M0	2	IB	J6	52	F	Invasive ductal carcinoma	T2N0M0	2	IB
D8	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J7	56	F	Invasive ductal carcinoma	T2N0M0	2	IB
D9	43	F	Invasive ductal carcinoma	T2N1M0	2	IB	J8	42	F	Invasive ductal carcinoma	T2N0M0	2	IB
D10	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J9	41	F	Invasive ductal carcinoma	T2N1M0	2	IB
D11	40	F	Invasive ductal carcinoma	T2N0M0	2	IA	J10	53	F	Invasive ductal carcinoma	T2N1M0	2	IB
D12	44	F	Invasive ductal carcinoma	T2N0M0	2	IA	J11	57	F	Invasive ductal carcinoma	T2N0M0	2	IB
D13	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	K1	45	F	Invasive ductal carcinoma	T2N0M0	2	IB
D14	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	K2	46	F	Invasive ductal carcinoma	T2N0M0	2	IB
D15	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	K3	48	F	Invasive ductal carcinoma	T2N1M0	2	IB
D16	48	F	Invasive ductal carcinoma	T2N0M0	2	IA	K4	48	F	Invasive ductal carcinoma	T2N1M0	2	IB

Fig. 175B

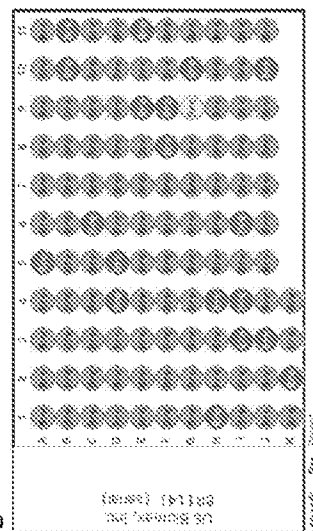


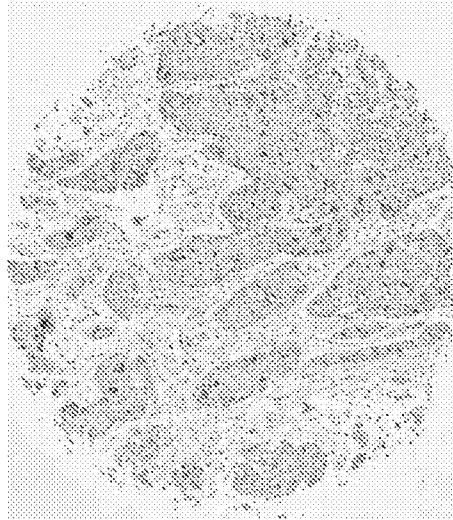
Figure 175A-175C

**N+20/C-27 antibody 32C1 IgG 0.25 ug/mL**  
**Breast cancer tissue array Br1141**

**Fig. 176A**

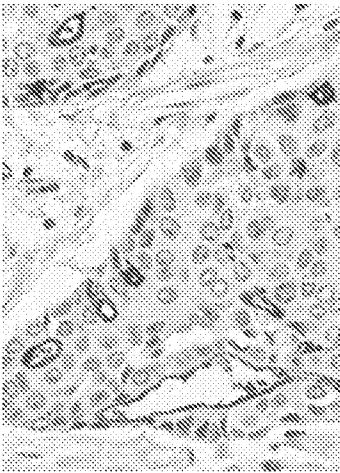
Position: B1

Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



**Fig. 176D**

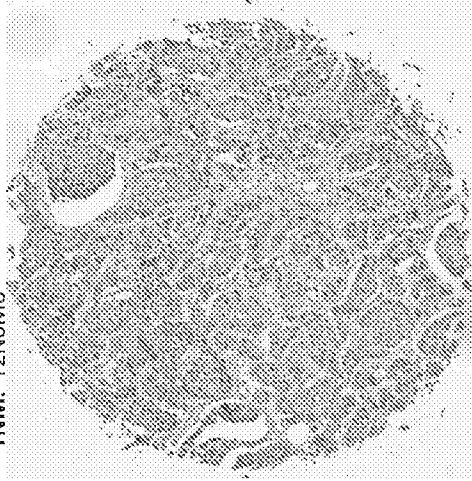
6x



**Fig. 176B**

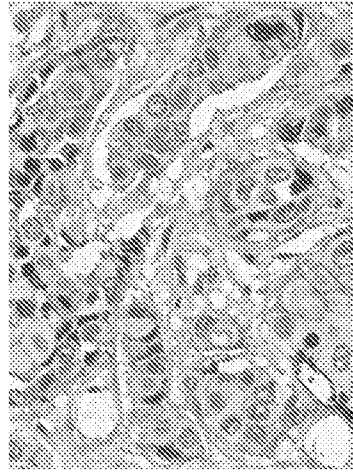
Position: B8

Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



**Fig. 176E**

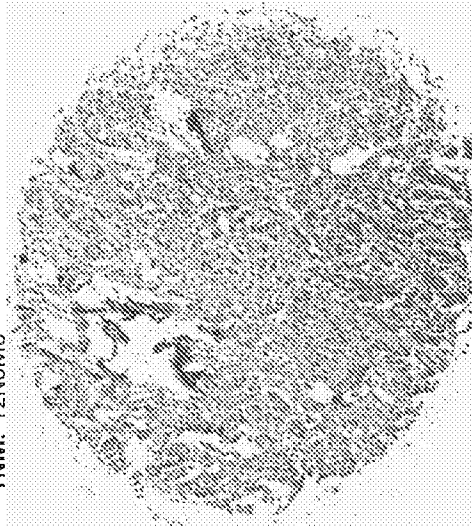
6x



**Fig. 176C**

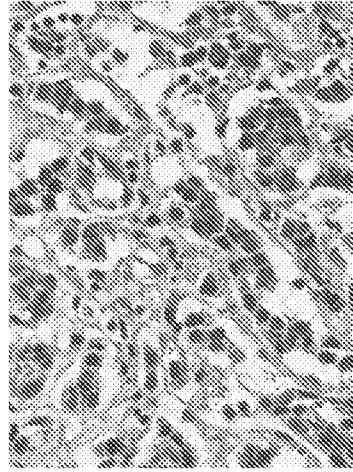
Position: G11

Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0



**Fig. 176F**

6x



**Fig. 176A-176F**

N+20/C-27 antibody 32C1 IgG 1-15 ug/mL  
Esophageal cancer tissue array ES1001

177A

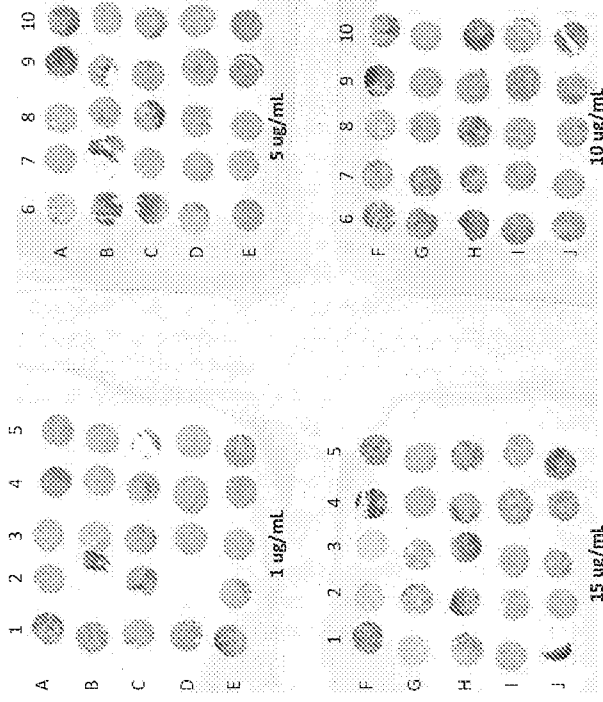


Fig. 1778

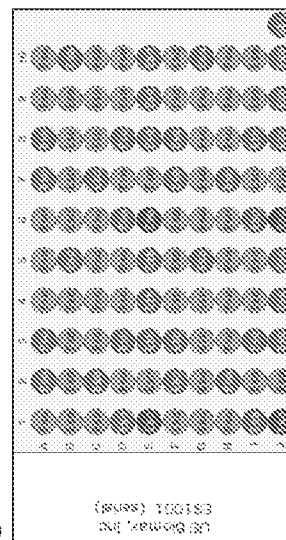


Fig. 177A-177C

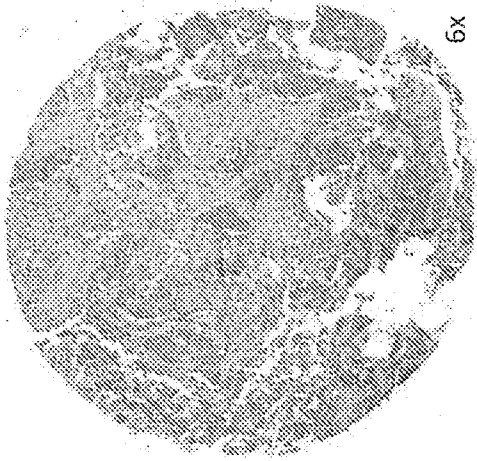
Ex. 177C

[illegible]

**N+20/C-27 antibody 32C1 IgG 1 ug/mL**  
**Esophageal cancer tissue array ES1001**

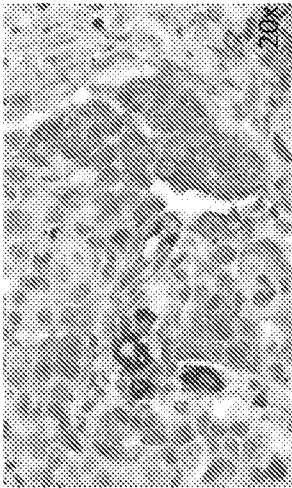
**Fig. 178A**

Position: D1  
Cell Type: Squamous cell carcinoma  
Tumor Grade: 2  
TNM: T3N1M0



6x

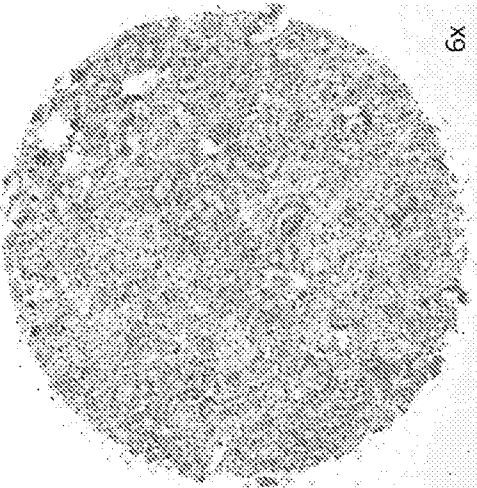
**Fig. 178D**



20x

**Fig. 178B**

Position: E4  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N1M0



6x

**Fig. 178E**



20x

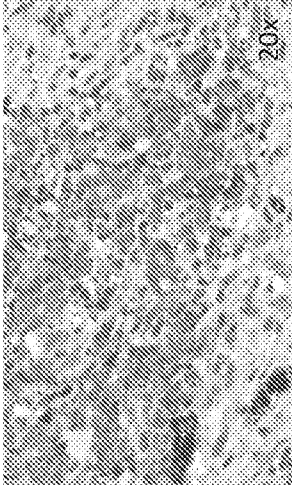
**Fig. 178C**

Position: E5  
Cell Type: Squamous cell carcinoma  
Tumor Grade: 3  
TNM: T3N1M0



6x

**Fig. 178F**



20x

**Fig. 178A-178F**

N+20/C-27 antibody 45C11 IgG 12.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 179A

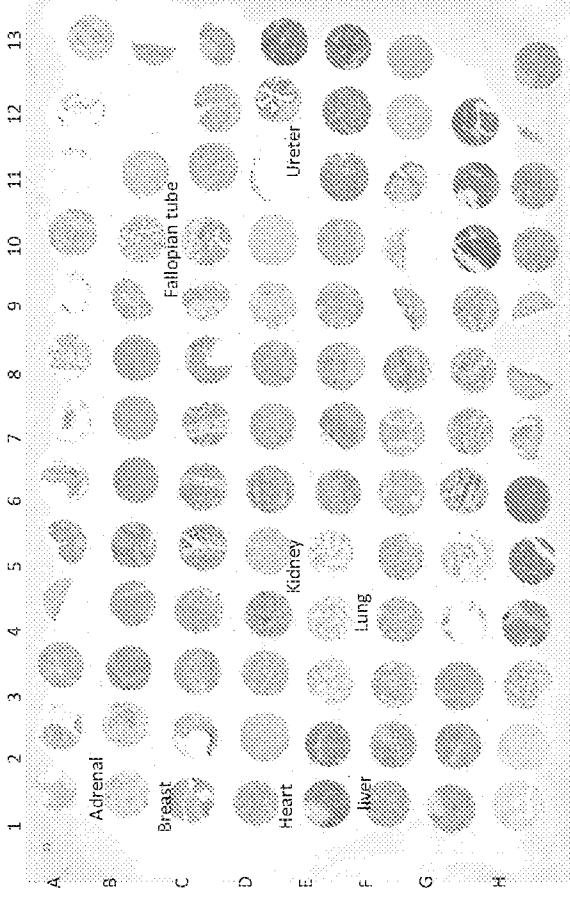


Fig. 179C

Position	Age	Sex	Organ/Anatomic Site	Pathology	Position	Age	Sex	Organ/Anatomic Site	Pathology
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	53	Adrenal gland	Normal	E3	F	72	Lung	Normal
A4	M	72	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	45	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bone, bone marrow	Normal	E7	F	46	Ovary	Normal
A8	F	43	Bone, bone marrow	Normal	E8	F	46	Ovary	Normal
A9	F	44	Head and neck, salivary gland	Normal	E9	M	42	Parotids	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	26	Parotids	Normal
A11	M	55	Eye	Normal	E11	F	59	Parotids	Normal
A12	M	50	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	F	35	Breast	Normal	E13	M	72	Parathyroid	Adenoma
B1	F	38	Breast	Normal	F1	F	3	Mammary gland	Normal
B2	F	33	Breast	Normal	F2	F	32	Mammary gland	Normal
B3	M	58	Brain, cerebellum	Normal	F3	F	30	Placenta	Normal
B4	F	55	Brain, cerebellum	Normal	F4	F	36	Placenta	Normal
B5	M	58	Brain, cerebral cortex	Normal	F5	M	64	Placenta	Normal
B6	M	35	Brain, cerebral cortex	Normal	F6	M	85	Prostate	Normal
B7	F	32	Brain, cerebral cortex	Normal	F7	M	85	Prostate	Normal
B8	F	42	Fallopian tube	Normal	F8	M	32	Prostate	Normal
B9	F	32	Fallopian tube	Normal	F9	F	32	Skin	Normal
B10	F	24	Fallopian tube	Normal	F10	F	26	Skin	Normal
B11	F	45	Esophagus	Normal	F11	M	26	Skin	Normal
B12	M	34	Esophagus	Normal	F12	M	58	Spinal cord	Normal
B13	M	45	Esophagus	Normal	G1	M	27	Spleen	Normal
C1	M	49	Stomach	Normal	G2	M	30	Spleen	Normal
C2	M	77	Stomach	Normal	G3	M	31	Spleen	Normal
C3	M	45	Intestine, small intestine	Normal	G4	F	90	Skeletal muscle	Normal
C4	M	75	Intestine, small intestine	Normal	G5	M	49	Skeletal muscle	Normal
C5	F	75	Intestine, small intestine	Normal	G6	M	70	Skeletal muscle	Normal
C6	F	75	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C7	M	2	Intestine, colon	Normal	G8	M	40	Testis	Normal
C8	M	46	Intestine, colon	Normal	G9	M	77	Testis	Normal
C9	M	74	Intestine, colon	Normal	G10	M	15	Thymus	Normal
C10	M	75	Intestine, rectum	Normal	G11	M	26	Thymus	Normal
C11	F	77	Intestine, rectum	Normal	G12	F	9	Thymus	Normal
C12	M	36	Intestine, rectum	Normal	G13	F	26	Thyroid	Normal
C13	M	38	Heart	Normal	H1	F	37	Thyroid	Normal
D1	M	35	Heart	Normal	H2	F	51	Thyroid	Normal
D2	F	34	Heart	Normal	H3	M	46	Tonsil	Normal
D3	F	52	Kidney, cortex	Normal	H4	M	37	Tonsil	Normal
D4	M	23	Kidney, cortex	Normal	H5	M	13	Tonsil	Normal
D5	F	52	Kidney, cortex	Normal	H6	F	38	Uterus, cervix	Normal
D6	M	53	Kidney, medulla	Schwannoma	H7	F	35	Uterus, cervix	Normal
D7	M	53	Kidney, medulla	Schwannoma	H8	F	44	Uterus, cervix	Normal
D8	M	39	Peripheral nerve	Schwannoma	H9	F	36	Uterus	Normal
D9	F	28	Peripheral nerve	Schwannoma	H10	F	36	Uterus	Normal
D10	F	44	Ureter	Normal	H11	F	41	Uterus	Normal
D11	F	42	Ureter	Normal	H12	F	46	Uterus	Normal
D12	F	57	Liver	Normal	H13	M	56	Skin	Malignant melanoma (tissue marker)

Fig. 179B

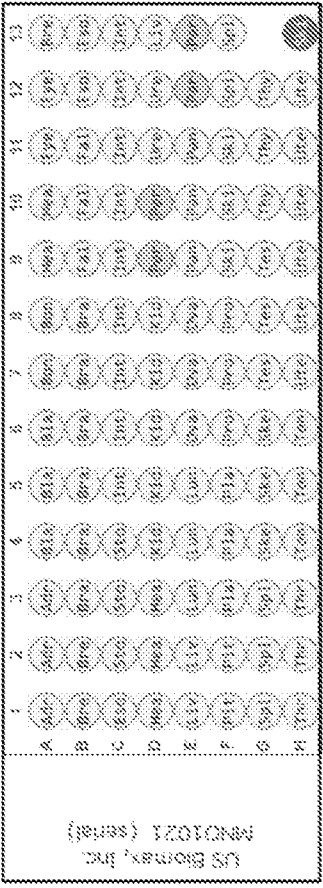
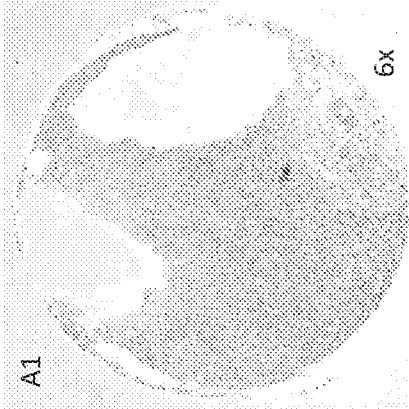


Figure 179A-179C

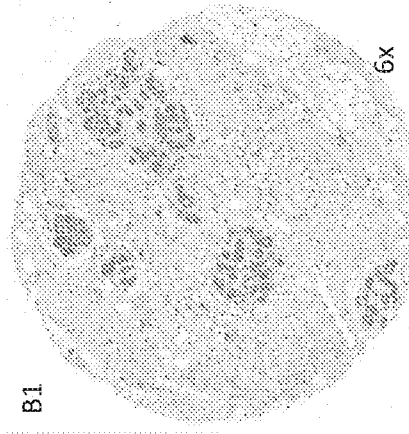


**N+20/C-27 antibody 45C11 IgG 12.5 ug/mL**  
**FDA Normal tissue array MNO1021**

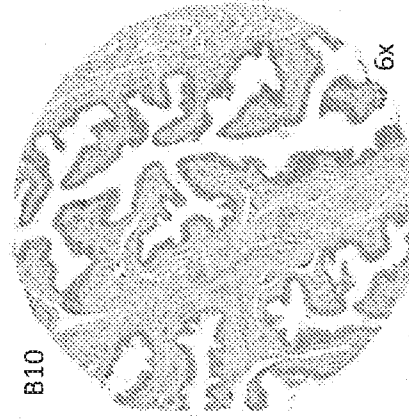
**Fig. 180A**  
**Normal Adrenal Gland**



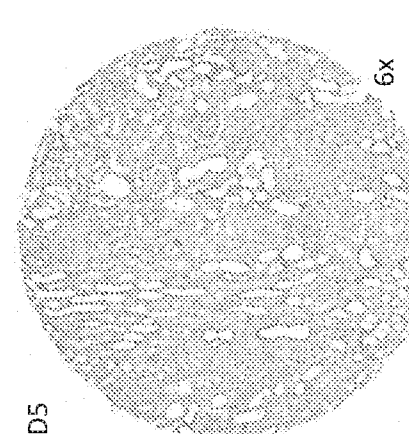
**Fig. 180B**  
**Normal Breast**



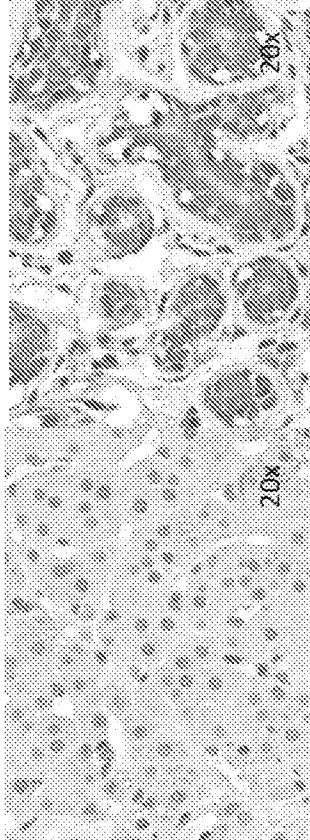
**Fig. 180C**  
**Normal Fallopian Tubes**



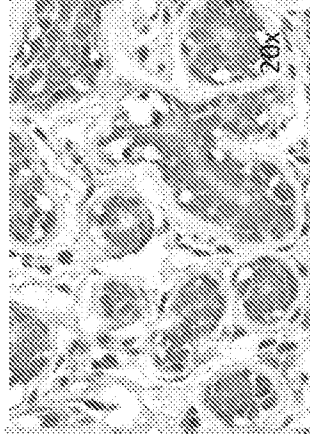
**Fig. 180D**  
**Normal Kidney**



**Fig. 180E**



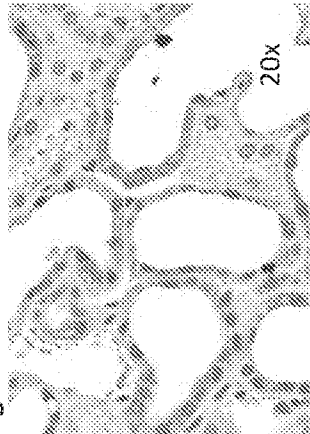
**Fig. 180F**



**Fig. 180G**



**Fig. 180H**



**Figure 180A-180H**

**N+20/C-27 antibody 45C11 IgG 12.5 ug/mL**  
**FDA Normal tissue array MNO1021**

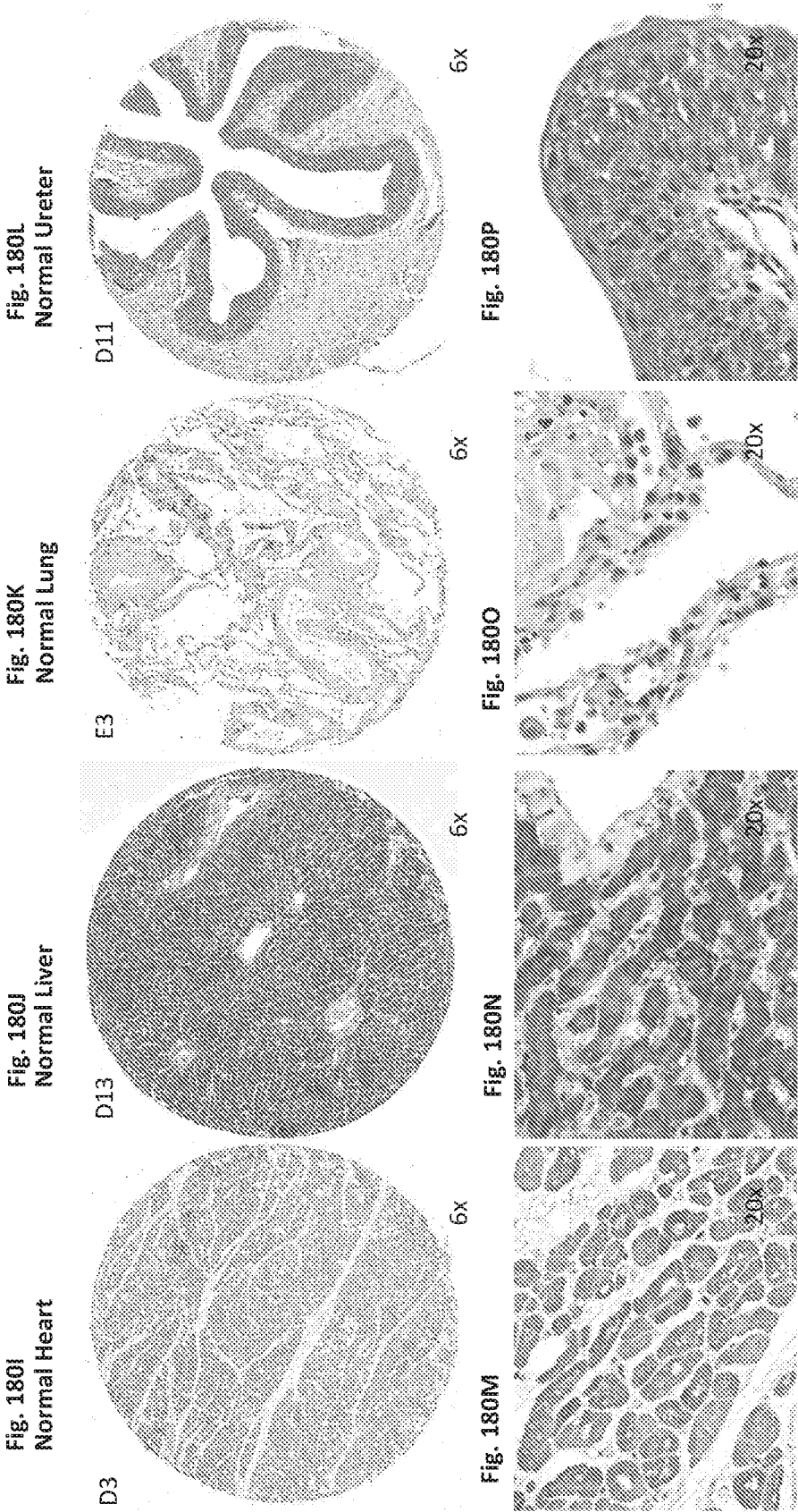


Figure 180I-180P



N+20/C-27 antibody 45C11 IgG 12.5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 180Q  
Normal eye



Fig. 180R  
Normal Cerebral cortex

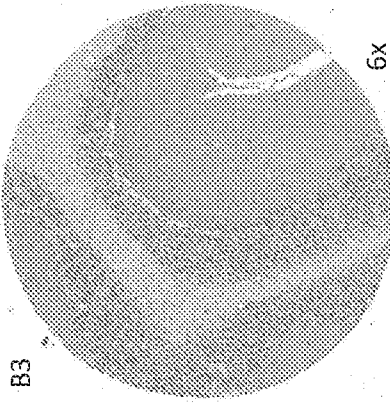


Fig. 180S  
Normal Bone marrow

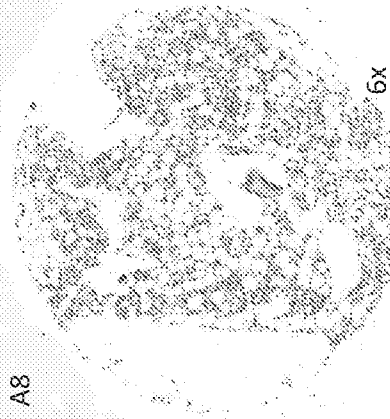


Fig. 180T  
Normal Skeletal muscle

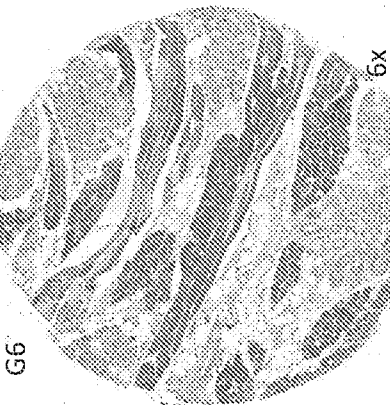


Fig. 180U



Fig. 180V

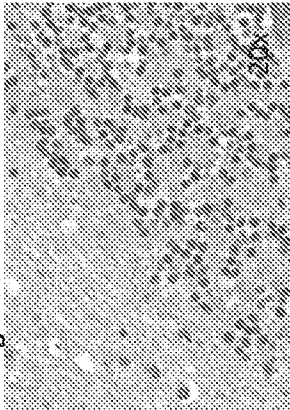


Fig. 180W



Fig. 180X

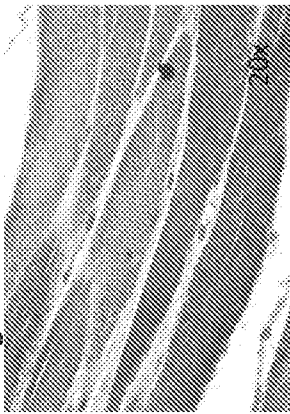


Figure 180Q-180X



N+20/C-27 antibody 45C11 IgG 10 ug/mL  
Breast cancer tissue array BR1007

Fig. 182A

Position: B4  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T1N1M0

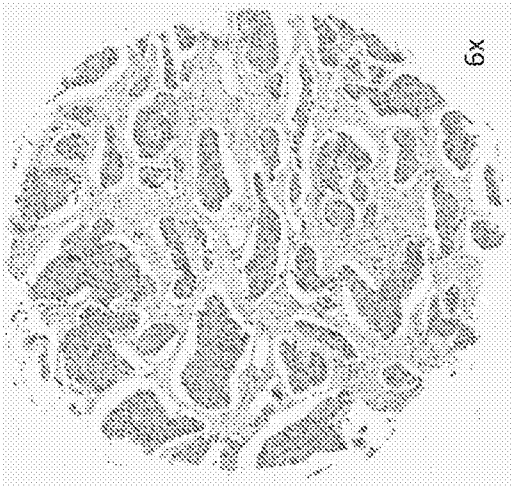


Fig. 182B

Position: B5  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N2M0

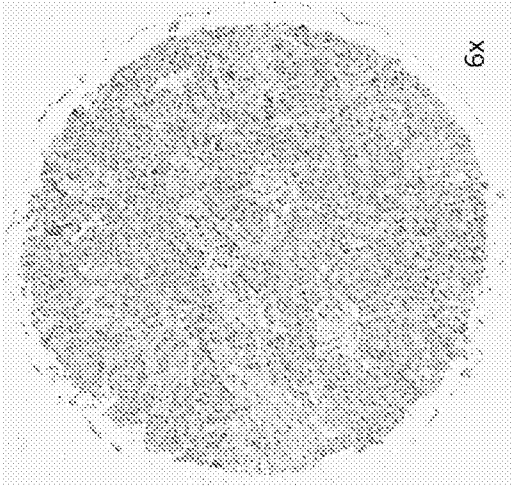


Fig. 182C

Position: D5  
Cell Type: Invasive Ductal Carcinoma  
Tumor Grade: 2  
TNM: T2N0M0

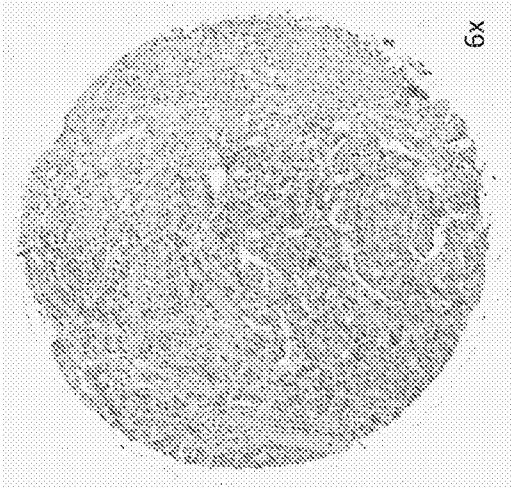


Fig. 182D

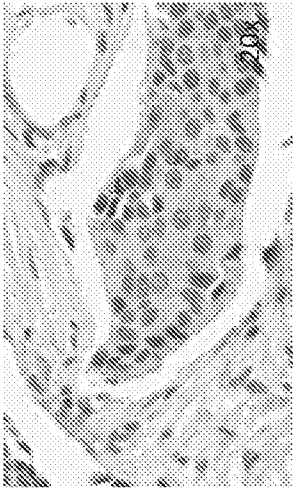


Fig. 182E

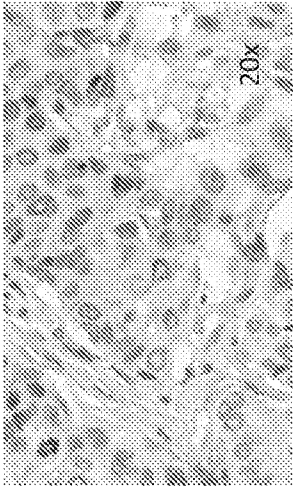


Fig. 182F



Fig. 182A-182F

N+20/C-27 antibody 45C11 5-15 ug/mL  
Pancreatic cancer tissue array PA1003

Ex. 183A

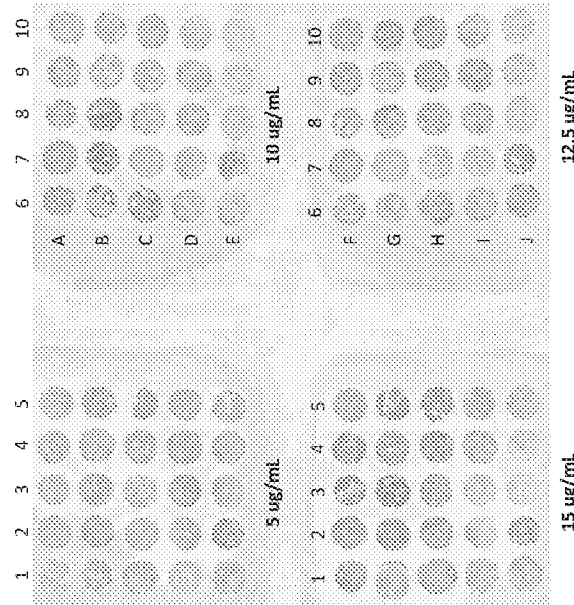
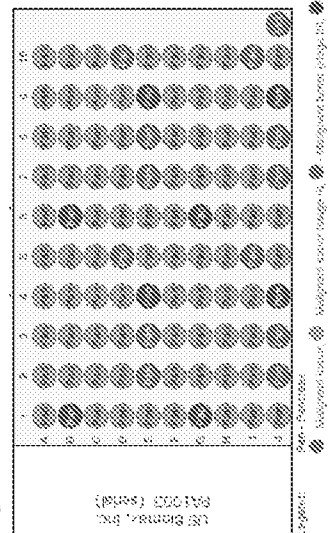
[illegible]

Fig. 183A-183C

3304

[illegible]

**N+20/C-27 antibody 45C11 12.5 ug/ml**  
**Pancreatic cancer tissue array PA805c**

**Fig. 184A**

Position: F6  
Cell Type: Invasive ductal carcinoma  
Tumor Grade: 2  
TNM: T3N1M0



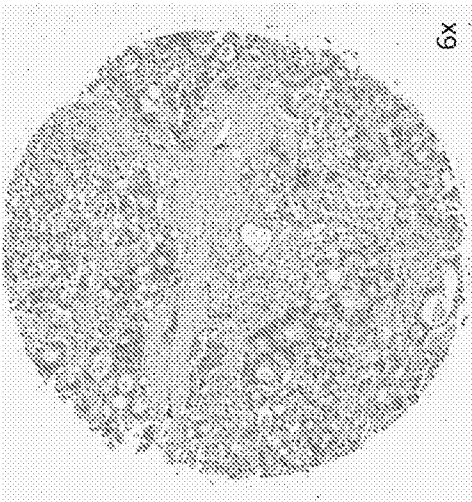
**Fig. 184B**

Position: H8  
Cell Type: Invasive ductal carcinoma  
Tumor Grade: 2  
TNM: T2N2M0

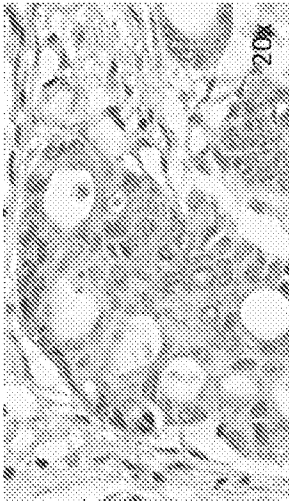


**Fig. 184C**

Position: I9  
Cell Type: Invasive ductal carcinoma  
Tumor Grade: 3  
TNM: T2N1M0



**Fig. 184D**



**Fig. 184E**



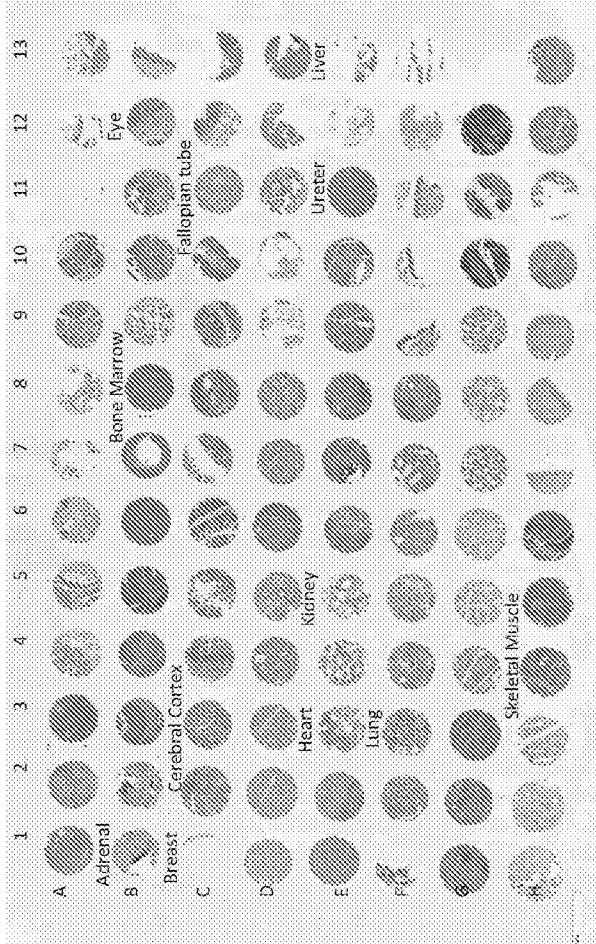
**Fig. 184F**



**Fig. 184A-184F**

**N+9/C-9 antibody 3C5 IgG 10 ug/mL**  
FDA Normal tissue array MNO1021

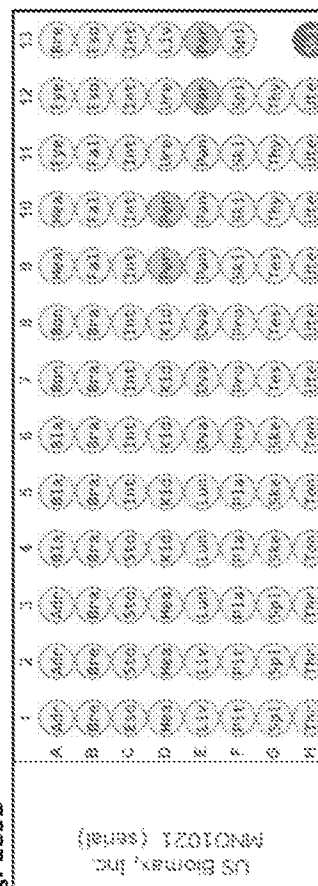
**Fig. 185A**



**Fig. 185C**

Position	Age	Sex	Organ/Anatomic Site	Pathology	Position	Age	Sex	Organ/Anatomic Site	Pathology
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	51	Adrenal gland	Normal	E3	F	72	Lung	Normal
A4	M	72	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	45	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bladder, urinary	Normal	E7	F	46	Ovary	Normal
A8	F	44	Bone, bone marrow	Normal	E8	F	46	Ovary	Normal
A9	F	44	Bone, bone marrow	Normal	E9	M	42	Prostate	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	26	Prostate	Normal
A11	M	55	Head and neck, salivary gland	Normal	E11	F	59	Prostate	Normal
A12	M	55	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	F	35	Eye	Normal	E13	M	72	Parathyroid	Adenoma
B1	F	38	Breast	Normal	F1	F	3	Mammary gland	Normal
B2	F	38	Breast	Normal	F2	F	32	Mammary gland	Normal
B3	M	58	Brain, cerebellum	Normal	F3	F	30	Placenta	Normal
B4	M	58	Brain, cerebellum	Normal	F4	F	27	Placenta	Normal
B5	M	58	Brain, cerebral cortex	Normal	F5	M	64	Prostate	Normal
B6	M	58	Brain, cerebral cortex	Normal	F6	M	65	Prostate	Normal
B7	M	58	Brain, cerebral cortex	Normal	F7	M	65	Prostate	Normal
B8	F	42	Fallopian tube	Normal	F8	F	32	Skin	Normal
B9	F	32	Fallopian tube	Normal	F9	F	32	Skin	Normal
B10	F	32	Fallopian tube	Normal	F10	M	26	Skin	Normal
B11	F	32	Fallopian tube	Normal	F11	M	26	Skin	Normal
B12	M	34	Esophagus	Normal	F12	M	58	Spinal cord	Normal
B13	M	34	Esophagus	Normal	G1	M	27	Spleen	Normal
C1	M	45	Esophagus	Normal	G2	M	30	Spleen	Normal
C2	M	45	Esophagus	Normal	G3	M	31	Spleen	Normal
C3	M	45	Esophagus	Normal	G4	F	60	Skeletal muscle	Normal
C4	M	45	Esophagus	Normal	G5	M	49	Skeletal muscle	Normal
C5	M	45	Esophagus	Normal	G6	M	70	Skeletal muscle	Normal
C6	F	75	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C7	F	75	Intestine, small intestine	Normal	G8	M	43	Testis	Normal
C8	M	2	Intestine, colon	Normal	G9	M	15	Thymus	Normal
C9	M	2	Intestine, colon	Normal	G10	M	15	Thymus	Normal
C10	M	2	Intestine, colon	Normal	G11	M	26	Thymus	Normal
C11	F	77	Intestine, rectum	Normal	G12	F	9	Thymus	Normal
C12	F	77	Intestine, rectum	Normal	G13	F	26	Thyroid	Normal
C13	M	36	Intestine, rectum	Normal	H1	F	37	Thyroid	Normal
D1	M	36	Intestine, rectum	Normal	H2	F	37	Thyroid	Normal
D2	M	36	Intestine, rectum	Normal	H3	F	51	Thyroid	Normal
D3	F	44	Heart	Normal	H4	M	46	Tonsil	Normal
D4	F	44	Heart	Normal	H5	M	37	Tonsil	Normal
D5	F	52	Kidney, cortex	Normal	H6	M	13	Tonsil	Normal
D6	F	52	Kidney, cortex	Normal	H7	F	36	Uterus, cervix	Normal
D7	M	53	Kidney, medulla	Normal	H8	F	35	Uterus, cervix	Normal
D8	M	53	Kidney, medulla	Schwannoma	H9	F	44	Uterus, cervix	Normal
D9	M	53	Kidney, medulla	Schwannoma	H10	F	36	Uterus	Normal
D10	F	29	Peripheral nerve	Schwannoma	H11	F	41	Uterus	Normal
D11	F	44	Peripheral nerve	Schwannoma	H12	F	46	Uterus	Normal
D12	F	44	Ureter	Normal	H13	M	56	Skin	Malignant melanoma (tissue marker)
D13	F	44	Ureter	Normal					

**Fig. 185B**



**Figure 185A-185C**



N+9/C-9 antibody 3C5 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 186A

Normal Adrenal Gland

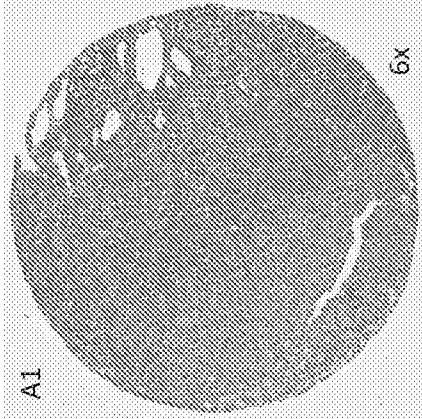


Fig. 186B

Normal Breast

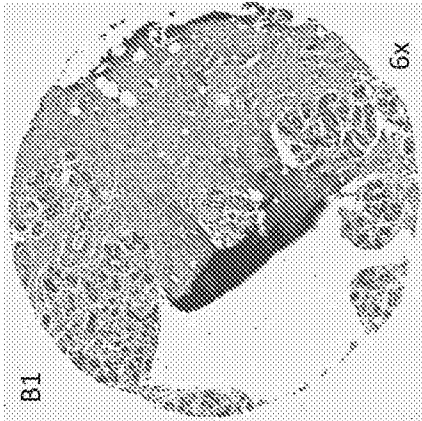


Fig. 186C

Normal Fallopian Tubes

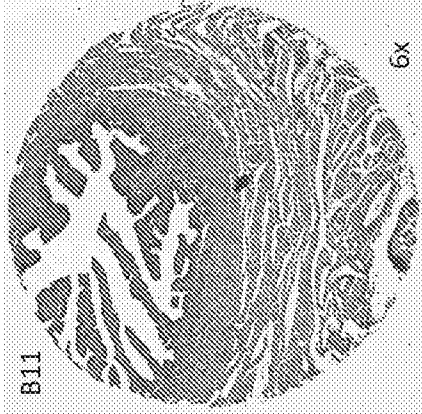


Fig. 186D

Normal Kidney

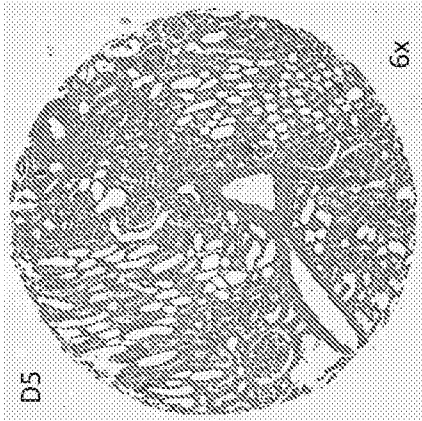


Fig. 186E

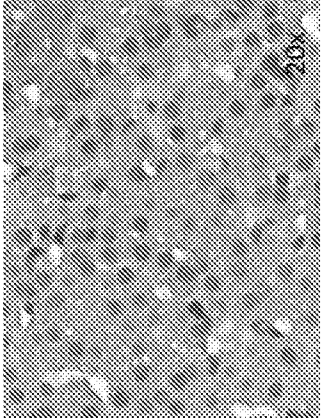


Fig. 186F

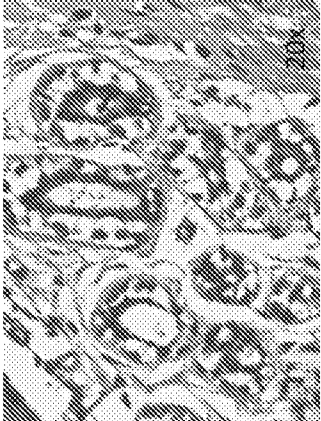


Fig. 186G



Fig. 186H

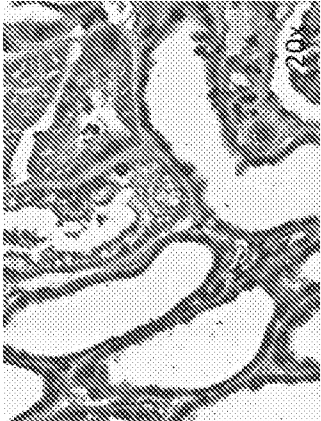


Figure 186A-186H

N+9/C-9 antibody 3C5 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 186I  
Normal Heart

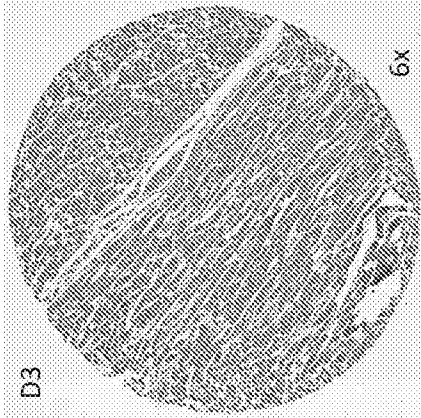


Fig. 186J  
Normal Liver

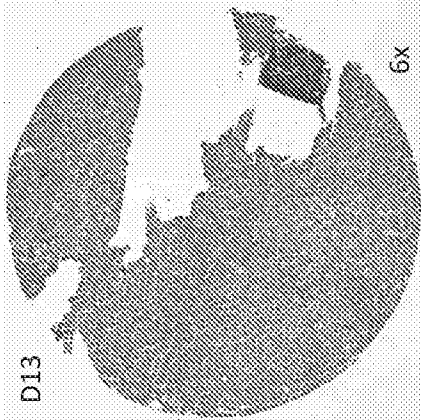


Fig. 186K  
Normal Lung

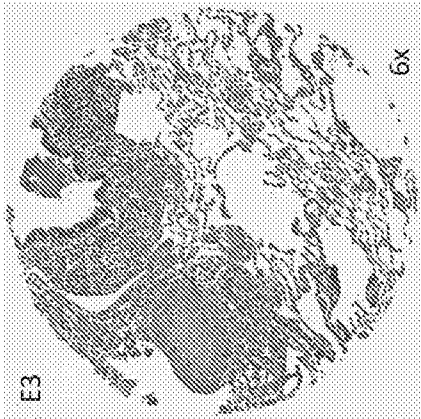


Fig. 186L  
Normal Ureter

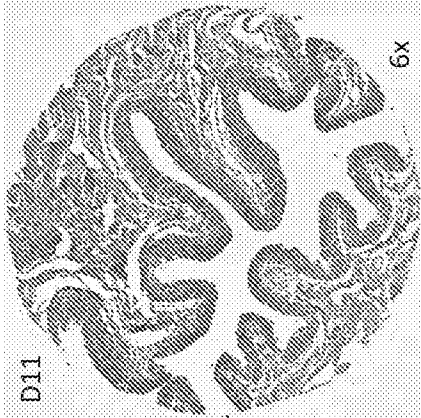


Fig. 186M

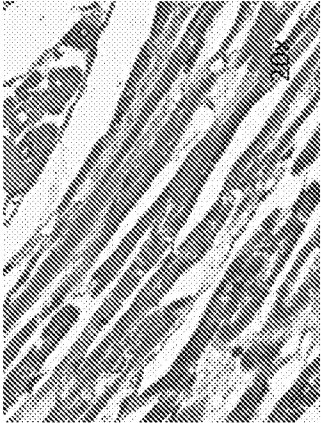


Fig. 186N

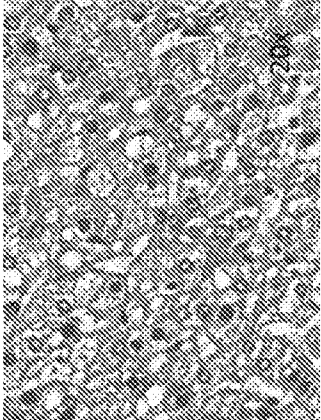


Fig. 186O

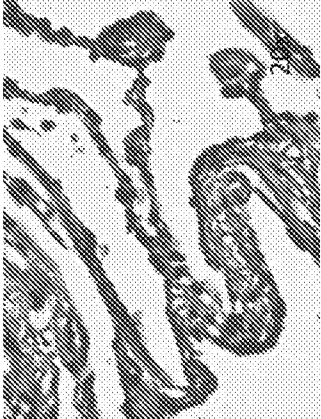


Fig. 186P

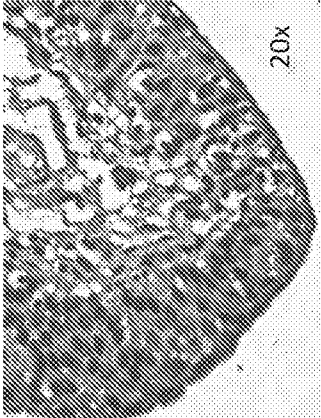


Figure 186I-186P



N+9/C-9 antibody 3C5 IgG 10 ug/mL  
FDA Normal tissue array MNO1021

Fig. 186Q  
Normal Eye

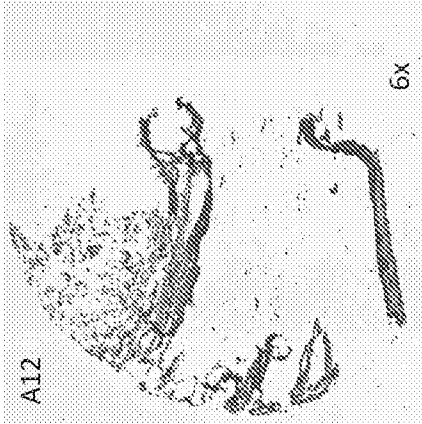


Fig. 186R  
Normal Cerebral Cortex

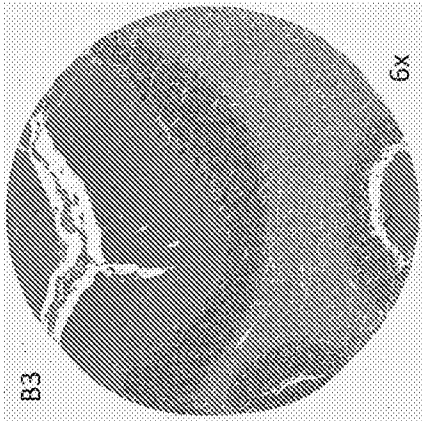


Fig. 186S  
Normal Bone Marrow

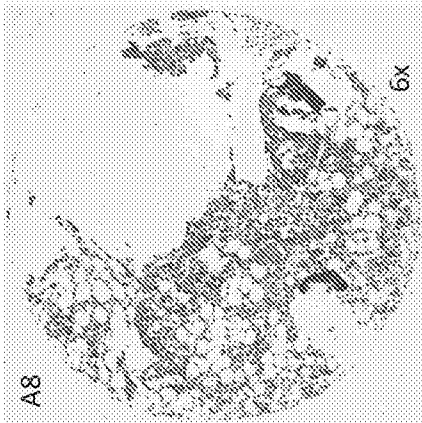


Fig. 186T  
Normal Skeletal Muscle

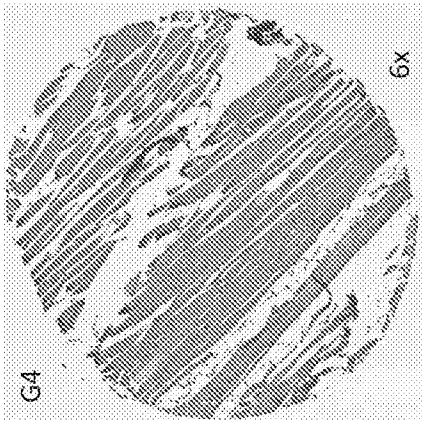


Fig. 186U



Fig. 186V

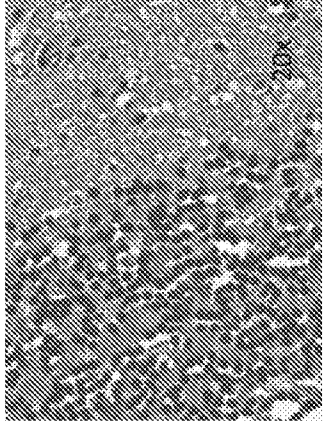


Fig. 186W

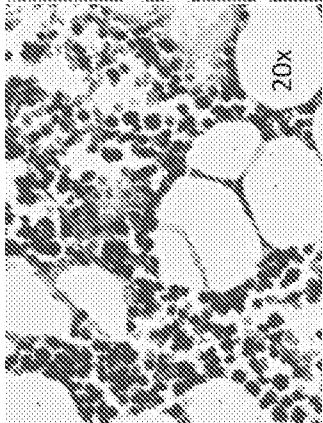


Fig. 186X

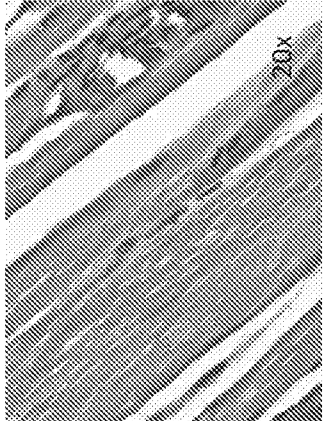
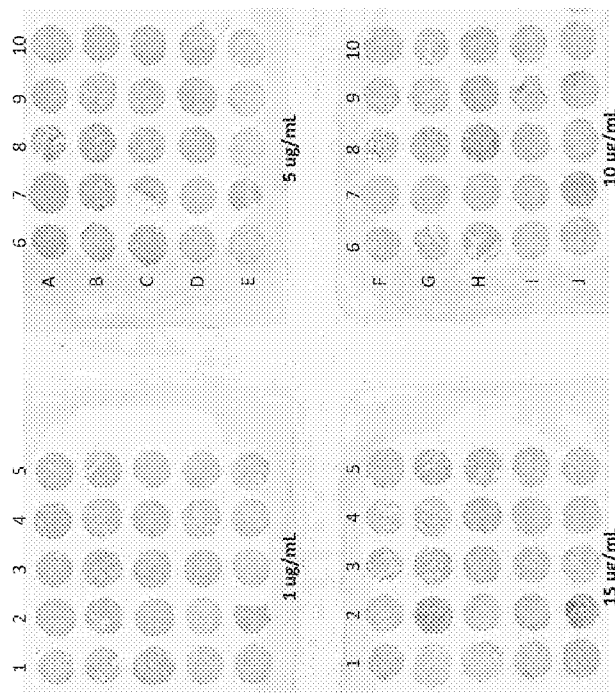


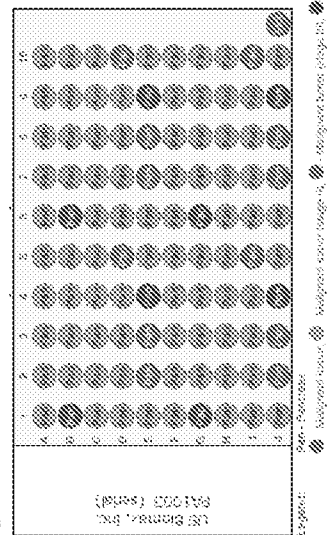
Figure 186Q-186X

**N+9/C-9 antibody 3C5 10 ug/mL**  
Pancreatic cancer tissue array PA1003

**Fig. 187A**



**Fig. 187B**



**Fig. 187A-187C**

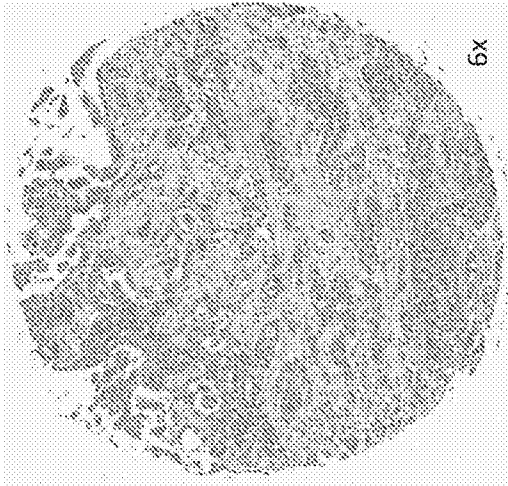
**Fig. 187C**

A1	38	F	Adenocarcinoma	TSNMO	2	II	71	35	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A2	41	M	Adenocarcinoma	TSNMO	2	II	52	44	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A3	59	M	Adenocarcinoma	TSNMO	2	II	53	35	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A4	65	M	Adenocarcinoma	TSNMO	2	II	54	65	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A5	53	M	Adenocarcinoma	TSNMO	2	II	55	53	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A6	39	F	Adenocarcinoma	TSNMO	2	II	56	35	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A7	44	M	Adenocarcinoma	TSNMO	2	II	57	44	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A8	59	M	Adenocarcinoma	TSNMO	2	II	58	59	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A9	65	M	Adenocarcinoma	TSNMO	2	II	59	65	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A10	53	M	Adenocarcinoma	TSNMO	2	II	60	53	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B1	42	M	Adenocarcinoma	TSNMO	2	II	61	42	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B2	52	M	Adenocarcinoma	TSNMO	2	II	62	52	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B3	57	M	Adenocarcinoma	TSNMO	2	II	63	57	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B4	34	M	Adenocarcinoma	TSNMO	2	II	64	34	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B5	71	F	Adenocarcinoma	TSNMO	2	II	65	71	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B6	42	M	Adenocarcinoma	TSNMO	2	II	66	42	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B7	52	M	Adenocarcinoma	TSNMO	2	II	67	52	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B8	57	M	Adenocarcinoma	TSNMO	2	II	68	57	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B9	34	M	Adenocarcinoma	TSNMO	2	II	69	34	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B10	72	F	Adenocarcinoma	TSNMO	2	II	70	72	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C1	47	M	Adenocarcinoma	TSNMO	2	II	71	47	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C2	64	F	Adenocarcinoma	TSNMO	2	II	72	64	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C3	51	M	Adenocarcinoma	TSNMO	2	II	73	51	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C4	68	F	Adenocarcinoma	TSNMO	2	II	74	68	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C5	41	M	Adenocarcinoma	TSNMO	2	II	75	41	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C6	47	M	Adenocarcinoma	TSNMO	2	II	76	47	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C7	64	F	Adenocarcinoma	TSNMO	2	II	77	64	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C8	51	M	Adenocarcinoma	TSNMO	2	II	78	51	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C9	68	F	Adenocarcinoma	TSNMO	2	II	79	68	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C10	41	M	Adenocarcinoma	TSNMO	2	II	80	41	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D1	58	F	Adenocarcinoma	TSNMO	2	II	81	58	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D2	41	F	Adenocarcinoma	TSNMO	2	II	82	41	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D3	77	M	Adenocarcinoma	TSNMO	2	II	83	77	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D4	59	M	Adenocarcinoma	TSNMO	2	II	84	59	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D5	58	F	Adenocarcinoma	TSNMO	2	II	85	58	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D6	72	F	Adenocarcinoma	TSNMO	2	II	86	72	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D7	41	F	Adenocarcinoma	TSNMO	2	II	87	41	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D8	72	F	Adenocarcinoma	TSNMO	2	II	88	72	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D9	59	M	Adenocarcinoma	TSNMO	2	II	89	59	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D10	59	M	Adenocarcinoma	TSNMO	2	II	90	59	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E1	53	F	Adenocarcinoma	TSNMO	2	II	91	53	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E2	60	M	Adenocarcinoma	TSNMO	2	II	92	60	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E3	56	M	Adenocarcinoma	TSNMO	2	II	93	56	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E4	40	F	Adenocarcinoma	TSNMO	2	II	94	40	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E5	56	F	Adenocarcinoma	TSNMO	2	II	95	56	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E6	53	F	Adenocarcinoma	TSNMO	2	II	96	53	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E7	60	M	Adenocarcinoma	TSNMO	2	II	97	60	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E8	56	M	Adenocarcinoma	TSNMO	2	II	98	56	M	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E9	60	F	Adenocarcinoma	TSNMO	2	II	99	60	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E10	66	F	Adenocarcinoma	TSNMO	2	II	100	66	F	Adenocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant

**N+9/C-9 antibody 3C5 10 ug/mL**  
**Pancreatic cancer tissue array PA1003**

**Fig. 188A**

Position: C3  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0

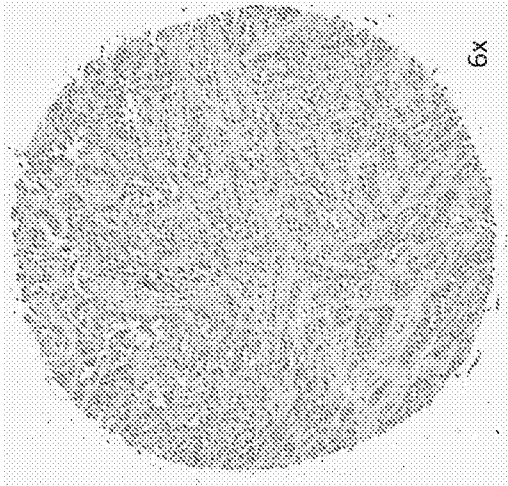


**Fig. 188D**



**Fig. 188B**

Position: C4  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0

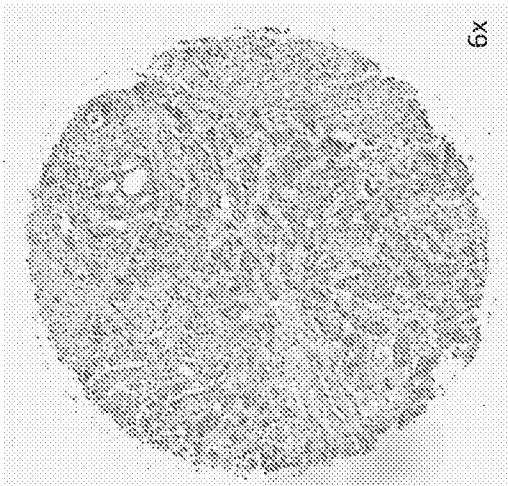


**Fig. 188E**

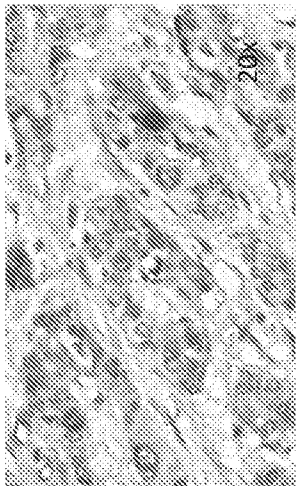


**Fig. 188C**

Position: E2  
Cell Type: Adenocarcinoma  
Tumor Grade: 2-3  
TNM: T3N1M0



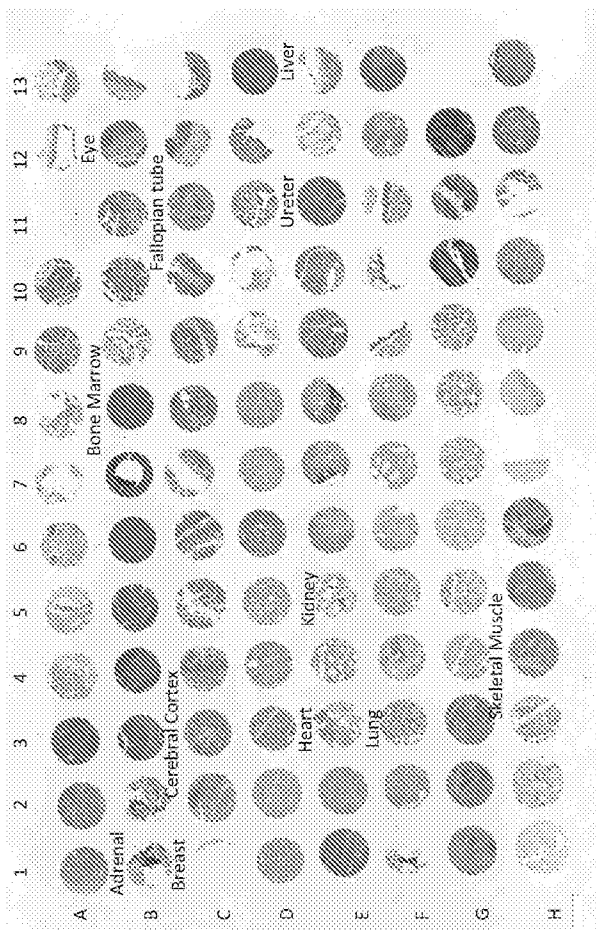
**Fig. 188F**



**Fig. 188A-188F**

**N+9/C-9 antibody 8A9 IgG 15 ug/mL**  
**FDA Normal tissue array MNO1021**

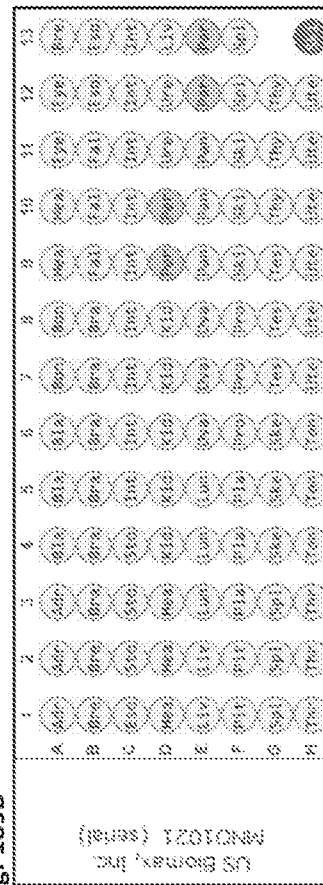
**Fig. 189A**



**Fig. 189C**

Position	Age	Sex	Organ/Anatomic Site	Pathology	Position	Age	Sex	Organ/Anatomic Site	Pathology
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	53	Adrenal gland	Normal	E3	M	72	Lung	Normal
A4	M	72	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	72	Bladder, urinary	Normal	E5	M	45	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bone, bone marrow	Normal	E7	F	46	Ovary	Normal
A8	F	44	Bone, bone marrow	Normal	E8	F	40	Ovary	Normal
A9	F	44	Head and neck, salivary gland	Normal	E9	M	42	Prostate	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	26	Prostate	Normal
A11	M	55	Eye	Normal	E11	M	59	Prostate	Normal
A12	M	50	Eye	Normal	E12	M	72	Parathyroid	Adenoma
A13	F	35	Breast	Normal	E13	M	72	Parathyroid	Adenoma
B1	F	35	Breast	Normal	F1	F	3	Mammary gland	Normal
B2	F	35	Breast	Normal	F2	F	32	Mammary gland	Normal
B3	M	58	Brain, cerebellum	Normal	F3	F	30	Placenta	Normal
B4	F	55	Brain, cerebellum	Normal	F4	F	27	Placenta	Normal
B5	M	58	Brain, cerebral cortex	Normal	F5	M	54	Prostate	Normal
B6	M	55	Brain, cerebral cortex	Normal	F6	M	55	Prostate	Normal
B7	F	42	Brain, cerebral cortex	Normal	F7	M	32	Prostate	Normal
B8	F	32	Fallopian tube	Normal	F8	F	32	Skin	Normal
B9	F	24	Fallopian tube	Normal	F9	M	26	Skin	Normal
B10	F	45	Esophagus	Normal	F10	M	26	Skin	Normal
B11	F	45	Esophagus	Normal	F11	M	58	Spinal cord	Normal
B12	M	34	Esophagus	Normal	F12	M	58	Spinal cord	Normal
B13	M	45	Esophagus	Normal	G1	M	27	Spleen	Normal
C1	M	45	Esophagus	Normal	G2	M	31	Spleen	Normal
C2	M	45	Esophagus	Normal	G3	M	31	Spleen	Normal
C3	M	45	Esophagus	Normal	G4	F	60	Skeletal muscle	Normal
C4	M	45	Esophagus	Normal	G5	M	49	Skeletal muscle	Normal
C5	M	45	Esophagus	Normal	G6	M	70	Skeletal muscle	Normal
C6	F	75	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C7	F	75	Intestine, small intestine	Normal	G8	M	40	Testis	Normal
C8	M	46	Intestine, colon	Normal	G9	M	15	Thymus	Normal
C9	M	46	Intestine, colon	Normal	G10	M	26	Thymus	Normal
C10	M	46	Intestine, colon	Normal	G11	M	26	Thymus	Normal
C11	F	77	Intestine, rectum	Normal	G12	F	9	Thymus	Normal
C12	F	77	Intestine, rectum	Normal	G13	F	26	Thyroid	Normal
C13	M	26	Intestine, rectum	Normal	H1	F	37	Thyroid	Normal
D1	M	45	Heart	Normal	H2	F	51	Thyroid	Normal
D2	M	45	Heart	Normal	H3	M	46	Tonsil	Normal
D3	F	44	Heart	Normal	H4	M	37	Tonsil	Normal
D4	F	52	Kidney, cortex	Normal	H5	M	13	Tonsil	Normal
D5	M	23	Kidney, cortex	Normal	H6	F	35	Uterus, cervix	Normal
D6	F	52	Kidney, medulla	Normal	H7	F	35	Uterus, cervix	Normal
D7	M	53	Kidney, medulla	Schwannoma	H8	F	44	Uterus, cervix	Normal
D8	M	53	Kidney, medulla	Schwannoma	H9	F	36	Uterus	Normal
D9	M	39	Peripheral nerve	Schwannoma	H10	F	36	Uterus	Normal
D10	F	28	Peripheral nerve	Schwannoma	H11	F	41	Uterus	Normal
D11	F	44	Ureter	Normal	H12	F	46	Uterus	Normal
D12	F	42	Ureter	Normal	H13	M	56	Skin	Malignant melanoma (tissue marker)
D13	F	57	Liver	Normal					

**Fig. 189B**



**Figure 189A-189C**

N+9/C-9 antibody 8A9 IgG 15 ug/mL  
FDA Normal tissue array MNO1021

Fig. 190A

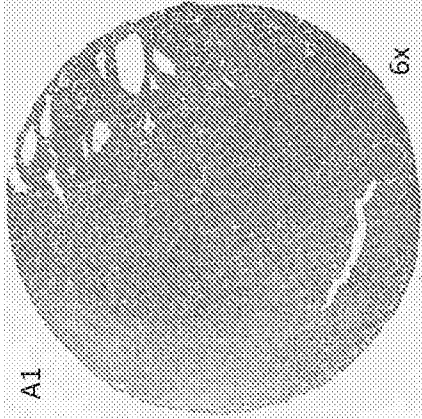


Fig. 190B

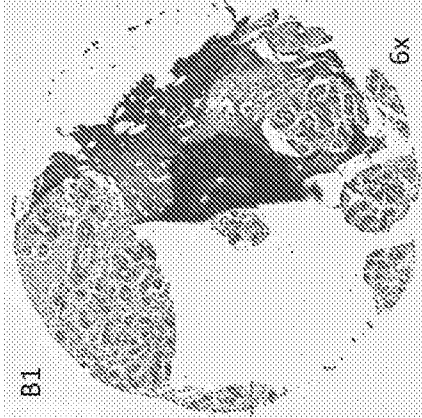


Fig. 190C

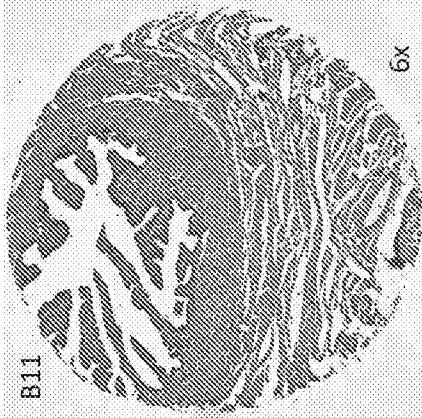


Fig. 190D

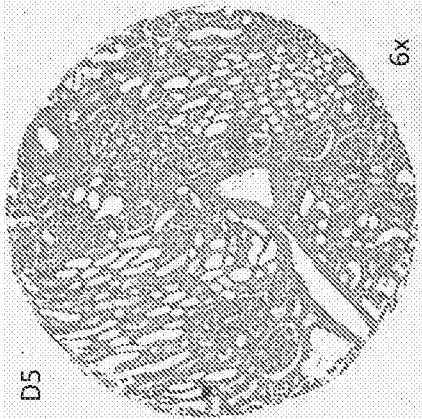


Fig. 190E

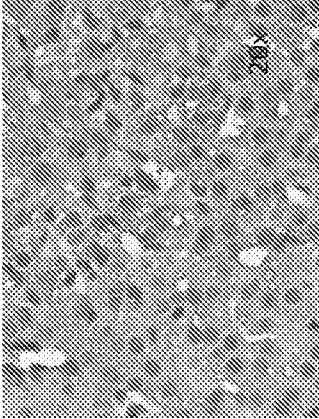


Fig. 190F

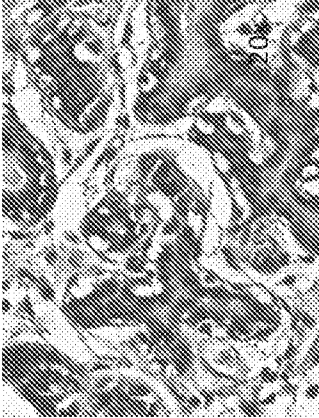


Fig. 190G

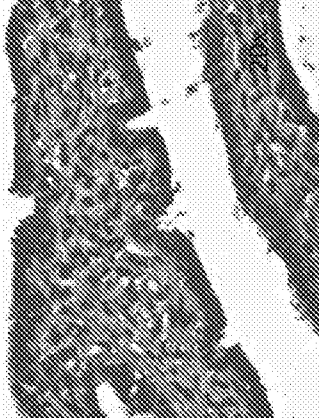


Fig. 190H

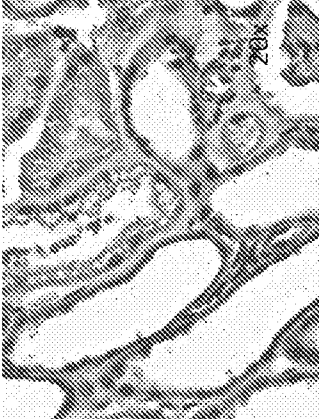
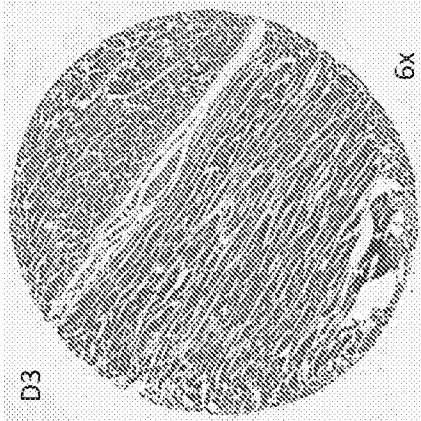


Figure 190A-190H

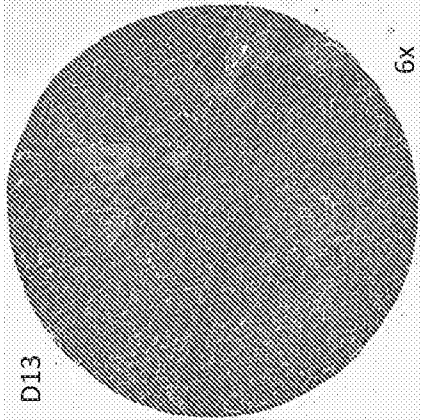


**N+9/C-9 antibody 8A9 IgG 15 ug/mL**  
**FDA Normal tissue array MNO1021**

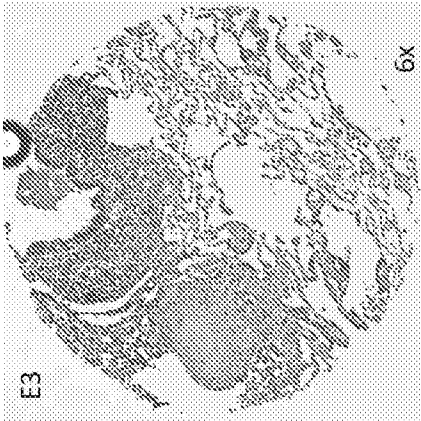
**Fig. 190I**  
**Normal Heart**



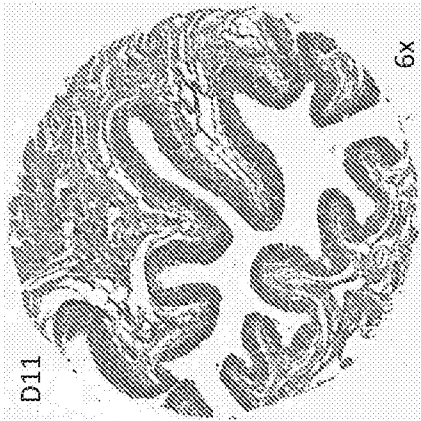
**Fig. 190J**  
**Normal Liver**



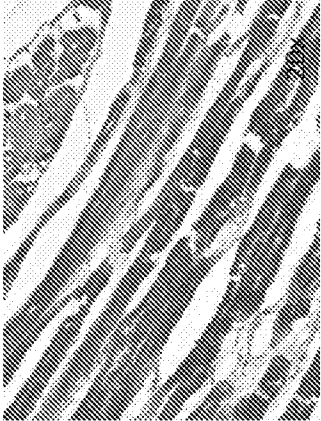
**Fig. 190K**  
**Normal Lung**



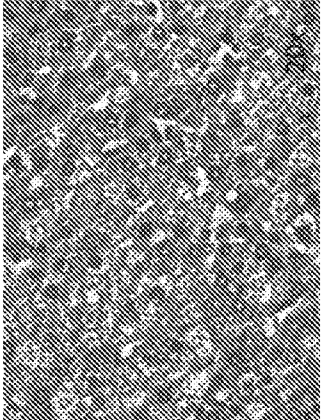
**Fig. 190L**  
**Normal Ureter**



**Fig. 190M**



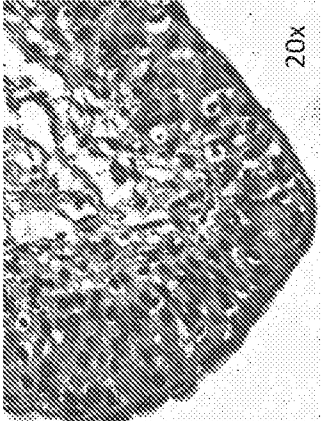
**Fig. 190N**



**Fig. 190O**



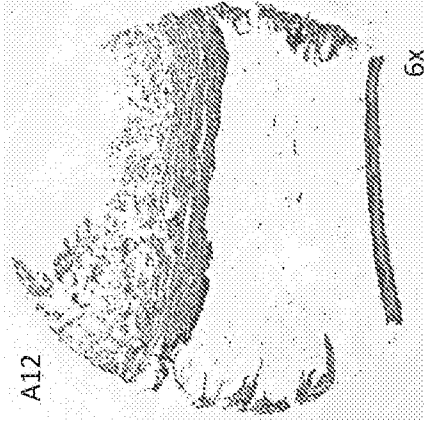
**Fig. 190P**



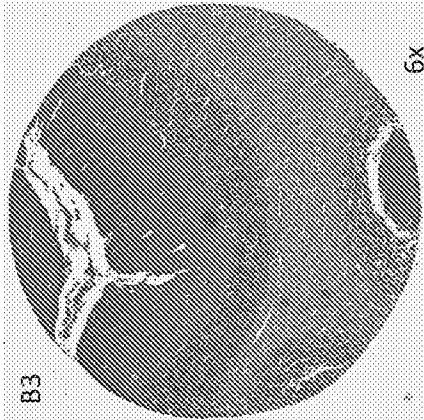
**Figure 190I-190P**

**N+9/C-9 antibody 8A9 IgG 15 ug/mL**  
**FDA Normal tissue array MNO1021**

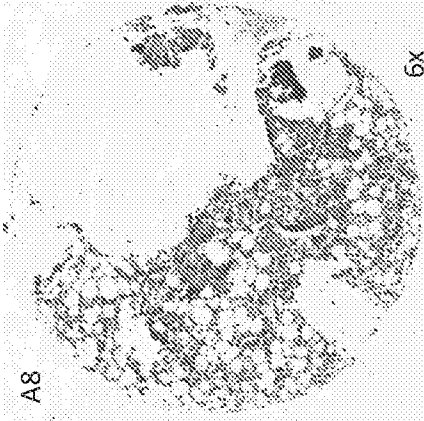
**Fig. 190Q**  
**Normal Eye**



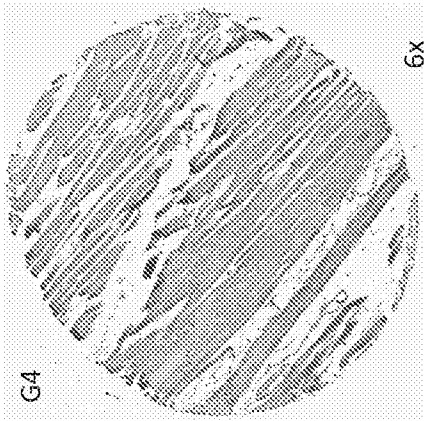
**Fig. 190R**  
**Normal Cerebral Cortex**



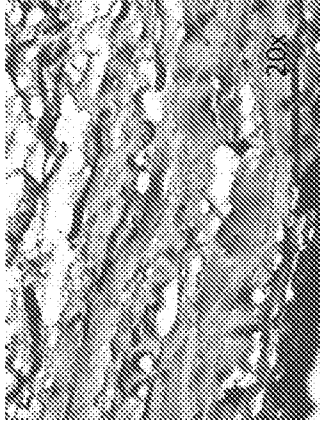
**Fig. 190S**  
**Normal Bone Marrow**



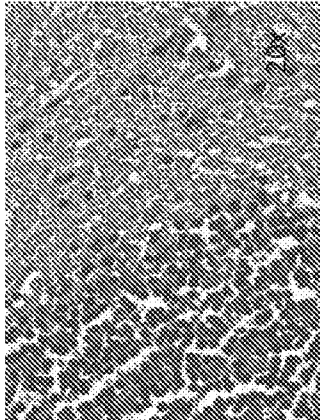
**Fig. 190T**  
**Normal Skeletal Muscle**



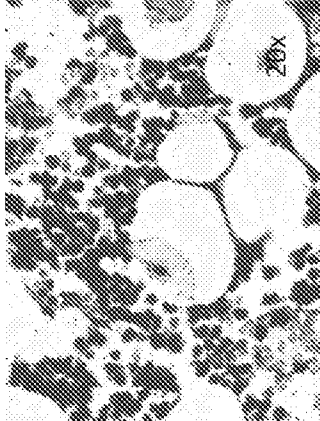
**Fig. 190U**



**Fig. 190V**



**Fig.**



**Fig. 190X**



**Figure 190Q-190X**

# N+9/C-9 antibody 8A9 15 ug/mL Pancreatic cancer tissue array PA1003

Fig. 191A

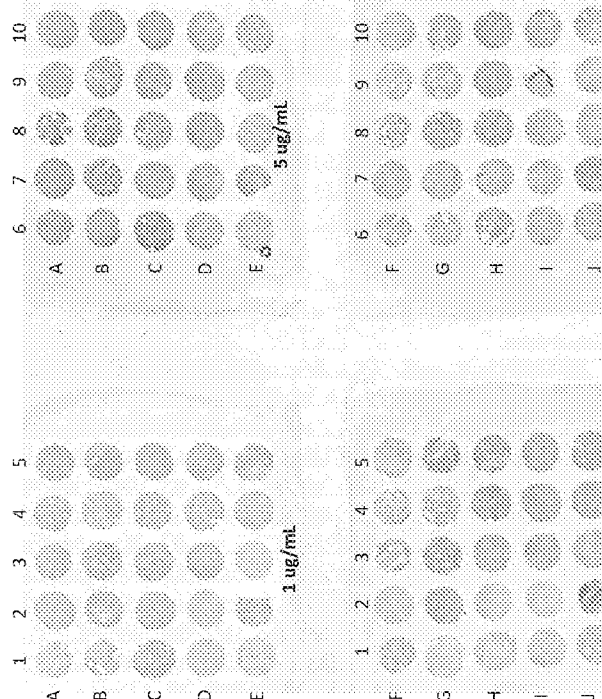
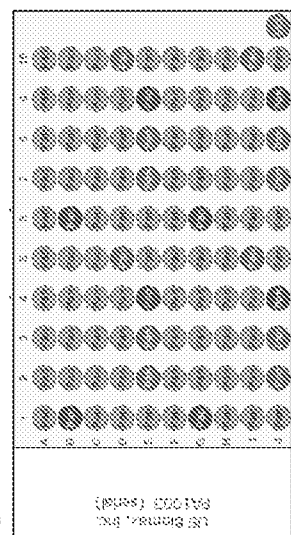


Fig. 191B



Legend:

8A9-Fluorescein, Adipose tissue (blue), Adipose tissue (blue), Adipose tissue (blue), Adipose tissue (blue), Adipose tissue (blue), Adipose tissue (blue), Adipose tissue (blue), Adipose tissue (blue), Adipose tissue (blue), Adipose tissue (blue)

Fig. 191A-191C

Fig. 191C

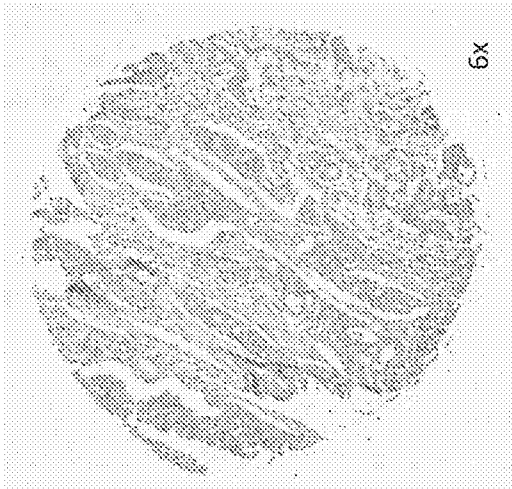
A1	8	F	Adipocarcinoma	TSNMO	2	II	71	35	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A2	41	M	Adipocarcinoma	TSNMO	2	II	52	44	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A3	59	M	Adipocarcinoma	TSNMO	2	II	53	35	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A4	65	M	Adipocarcinoma	TSNMO	2	II	54	65	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A5	53	M	Adipocarcinoma	TSNMO	2	II	55	53	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A6	39	F	Adipocarcinoma	TSNMO	2	II	56	35	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A7	44	M	Adipocarcinoma	TSNMO	2	II	57	44	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A8	55	M	Adipocarcinoma	TSNMO	2	II	58	55	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A9	65	M	Adipocarcinoma	TSNMO	2	II	59	65	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
A10	53	M	Adipocarcinoma	TSNMO	2	II	60	53	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B1	42	M	Adipocarcinoma	TSNMO	2	IV	61	42	M	Adipocarcinoma	TSNMO	2	IV	malignant	TSNMO	2	IV	malignant
B2	52	M	Adipocarcinoma	TSNMO	2	II	62	52	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B3	57	M	Adipocarcinoma	TSNMO	2	II	63	57	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B4	34	M	Adipocarcinoma	TSNMO	2	II	64	34	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B5	71	F	Adipocarcinoma	TSNMO	2	II	65	71	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B6	42	M	Adipocarcinoma	TSNMO	2	IV	66	42	M	Adipocarcinoma	TSNMO	2	IV	malignant	TSNMO	2	IV	malignant
B7	52	M	Adipocarcinoma	TSNMO	2	II	67	52	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B8	57	M	Adipocarcinoma	TSNMO	2	II	68	57	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B9	34	M	Adipocarcinoma	TSNMO	2	II	69	34	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
B10	72	F	Adipocarcinoma	TSNMO	2	II	70	72	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C1	47	M	Adipocarcinoma	TSNMO	2	II	71	47	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C2	64	F	Adipocarcinoma	TSNMO	2	II	72	64	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C3	51	M	Adipocarcinoma	TSNMO	2	II	73	51	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C4	68	F	Adipocarcinoma	TSNMO	2	II	74	68	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C5	41	M	Adipocarcinoma	TSNMO	2	II	75	41	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C6	47	M	Adipocarcinoma	TSNMO	2	II	76	47	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C7	64	F	Adipocarcinoma	TSNMO	2	II	77	64	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C8	51	M	Adipocarcinoma	TSNMO	2	II	78	51	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C9	68	F	Adipocarcinoma	TSNMO	2	II	79	68	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
C10	41	M	Adipocarcinoma	TSNMO	2	II	80	41	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D1	58	F	Adipocarcinoma	TSNMO	2	II	81	58	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D2	41	F	Adipocarcinoma	TSNMO	2	II	82	41	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D3	71	F	Adipocarcinoma	TSNMO	2	II	83	71	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D4	56	M	Adipocarcinoma	TSNMO	2	II	84	56	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D5	56	M	Adipocarcinoma	TSNMO	2	II	85	56	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D6	56	M	Adipocarcinoma	TSNMO	2	II	86	56	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D7	72	F	Adipocarcinoma	TSNMO	2	II	87	72	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D8	72	F	Adipocarcinoma	TSNMO	2	II	88	72	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D9	53	M	Adipocarcinoma	TSNMO	2	II	89	53	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
D10	53	M	Adipocarcinoma	TSNMO	2	II	90	53	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E1	53	M	Adipocarcinoma	TSNMO	2	II	91	53	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E2	60	M	Adipocarcinoma	TSNMO	2	II	92	60	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E3	56	M	Adipocarcinoma	TSNMO	2	II	93	56	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E4	46	F	Adipocarcinoma	TSNMO	2	II	94	46	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E5	56	M	Adipocarcinoma	TSNMO	2	II	95	56	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E6	56	M	Adipocarcinoma	TSNMO	2	II	96	56	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E7	60	M	Adipocarcinoma	TSNMO	2	II	97	60	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E8	56	M	Adipocarcinoma	TSNMO	2	II	98	56	M	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant
E9	60	F	Adipocarcinoma	TSNMO	2	IV	99	60	F	Adipocarcinoma	TSNMO	2	IV	malignant	TSNMO	2	IV	malignant
E10	66	F	Adipocarcinoma	TSNMO	2	II	100	66	F	Adipocarcinoma	TSNMO	2	II	malignant	TSNMO	2	II	malignant



**N+9/C-9 antibody 8A9 15 ug/mL**  
**Pancreatic cancer tissue array PA1003**

**Fig. 192A**

Position: A1  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0

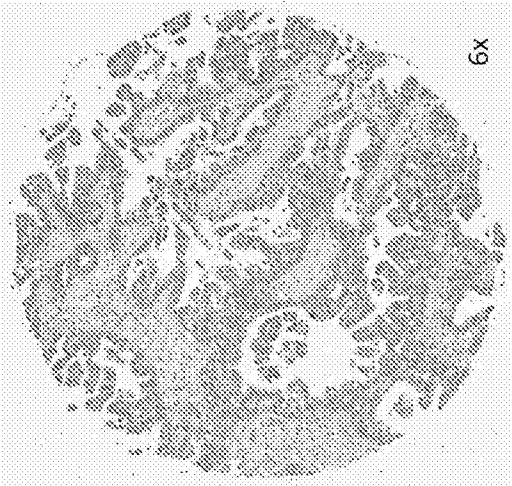


**Fig. 192D**



**Fig. 192B**

Position: B3  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0

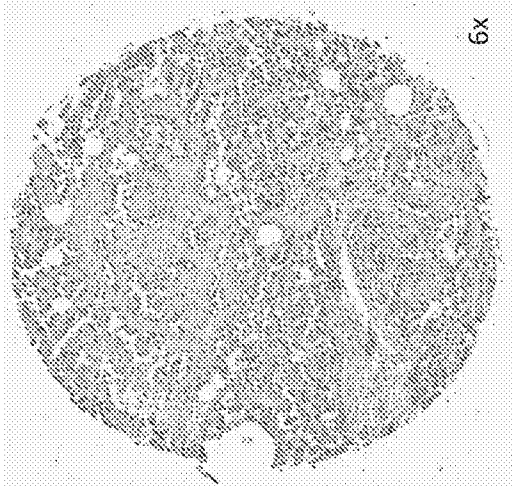


**Fig. 192E**



**Fig. 192C**

Position: C4  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



**Fig. 192F**



**Fig. 192A-192F**

N+9/C-9 antibody 17H6 30 ug/mL  
FDA Normal tissue array MNO1021

Fig. 193A

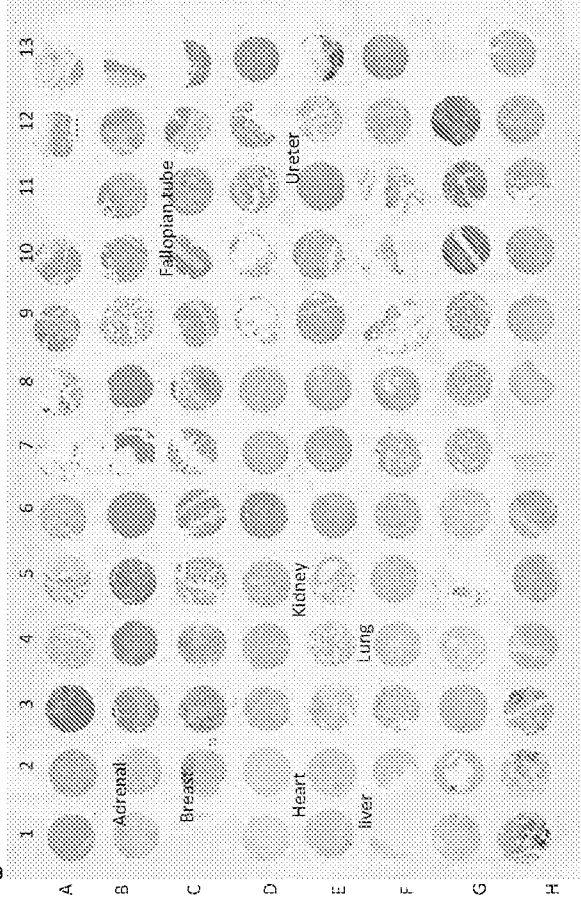


Fig. 193B

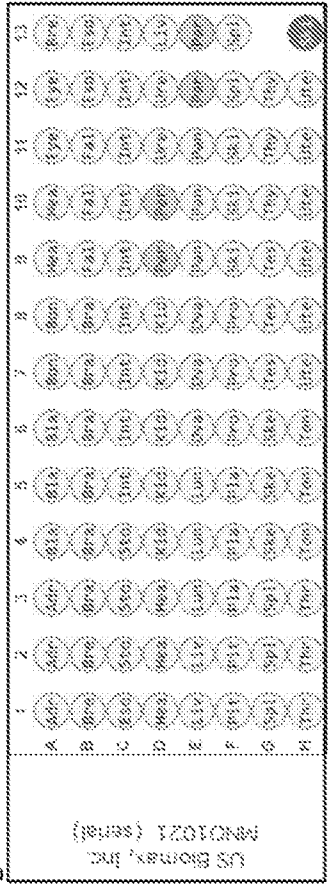


Figure 193A-193C

Fig. 193C

Position	Age	Sex	Organ/Anatomic Site	Pathology diagnosis	Position	Age	Sex	Organ/Anatomic Site	Pathology diagnosis
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	53	Adrenal gland	Normal	E3	M	72	Liver	Normal
A4	M	75	Bladder, urinary	Normal	E4	F	55	Lung	Normal
A5	M	75	Bladder, urinary	Normal	E5	M	43	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bone marrow	Normal	E7	F	46	Ovary	Normal
A8	F	44	Bone marrow	Normal	E8	F	40	Ovary	Normal
A9	F	44	Bone marrow	Normal	E9	M	42	Pancreas	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	35	Pancreas	Normal
A11	M	55	Head and neck, salivary gland	Normal	E11	F	59	Pancreas	Normal
A12	M	95	Eye	Normal	E12	M	17	Parathyroid	Adenoma
A13	M	35	Breast	Normal	E13	M	72	Parathyroid	Adenoma
B1	F	38	Breast	Normal	F1	F	3	Pituitary gland	Normal
B2	F	38	Breast	Normal	F2	F	32	Pituitary gland	Normal
B3	M	55	Brain, cerebellum	Normal	F3	F	30	Placenta	Normal
B4	F	55	Brain, cerebellum	Normal	F4	F	30	Placenta	Normal
B5	M	58	Brain, cerebral cortex	Normal	F5	F	64	Prostate	Normal
B6	M	85	Brain, cerebral cortex	Normal	F6	M	85	Prostate	Normal
B7	F	42	Fallopian tube	Normal	F7	M	85	Prostate	Normal
B8	F	32	Fallopian tube	Normal	F8	F	32	Shin	Normal
B9	F	32	Fallopian tube	Normal	F9	F	32	Shin	Normal
B10	M	24	Fallopian tube	Normal	F10	M	25	Shin	Normal
B11	M	45	Esophagus	Normal	F11	M	47	Spinal cord	Normal
B12	F	38	Esophagus	Normal	F12	M	56	Spinal cord	Normal
B13	F	38	Esophagus	Normal	G1	M	27	Spleen	Normal
C1	M	49	Stomach	Normal	G2	M	90	Spleen	Normal
C2	M	49	Stomach	Normal	G3	M	91	Spleen	Normal
C3	M	77	Stomach	Normal	G4	F	49	Skeletal muscle	Normal
C4	M	45	Intestine, small intestine	Normal	G5	F	70	Skeletal muscle	Normal
C5	F	75	Intestine, small intestine	Normal	G6	M	43	Testis	Normal
C6	F	75	Intestine, small intestine	Normal	G7	M	43	Testis	Normal
C7	M	2	Intestine, small intestine	Normal	G8	M	43	Testis	Normal
C8	M	48	Intestine, colon	Normal	G9	M	77	Thyroid	Normal
C9	M	48	Intestine, colon	Normal	G10	M	15	Thyroid	Normal
C10	M	74	Intestine, rectum	Normal	G11	M	28	Thyroid	Normal
C11	F	47	Intestine, rectum	Normal	G12	F	9	Thyroid	Normal
C12	F	75	Intestine, rectum	Normal	G13	F	26	Thyroid	Normal
C13	M	86	Intestine, rectum	Normal	H1	F	26	Thyroid	Normal
D1	M	38	Heart	Normal	H2	F	57	Thyroid	Normal
D2	M	38	Heart	Normal	H3	F	51	Thyroid	Normal
D3	F	34	Heart	Normal	H4	M	46	Tonsil	Normal
D4	F	32	Kidney, cortex	Normal	H5	M	37	Tonsil	Normal
D5	M	23	Kidney, cortex	Normal	H6	M	13	Tonsil	Normal
D6	M	52	Kidney, cortex	Normal	H7	F	38	Uterus, cervix	Normal
D7	M	52	Kidney, medulla	Normal	H8	F	35	Uterus, cervix	Normal
D8	M	23	Kidney, medulla	Schwannoma	H9	F	44	Uterus, cervix	Normal
D9	M	39	Peripheral nerve	Schwannoma	H10	F	38	Uterus, endometrium	Normal
D10	F	29	Peripheral nerve	Schwannoma	H11	F	41	Uterus, endometrium	Normal
D11	F	44	Ureter	Normal	H12	F	46	Uterus, endometrium	Normal
D12	F	42	Ureter	Normal	H13	M	58	Skin	Malignant melanoma (tissue mass)
D13	F	57	Liver	Normal					

N+9/C-9 antibody 17H6 30 ug/mL  
FDA Normal tissue array MNO1021

Fig. 194A  
Normal Adrenal Gland

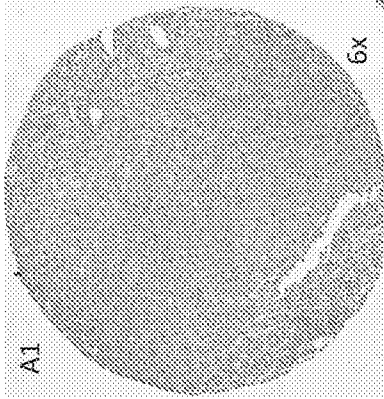


Fig. 194B  
Normal Breast

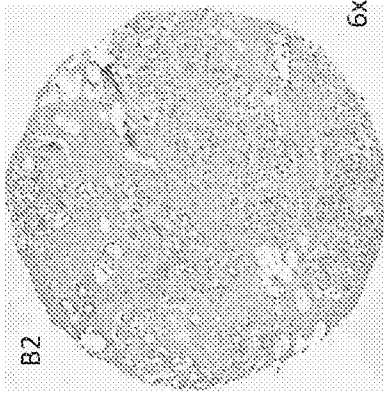


Fig. 194C  
Normal Fallopian Tubes

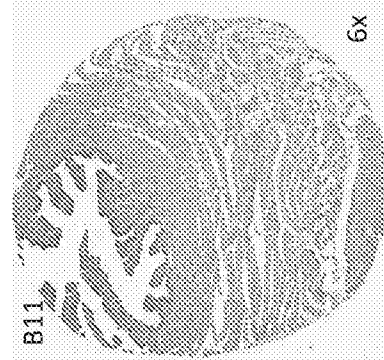


Fig. 194D  
Normal Kidney

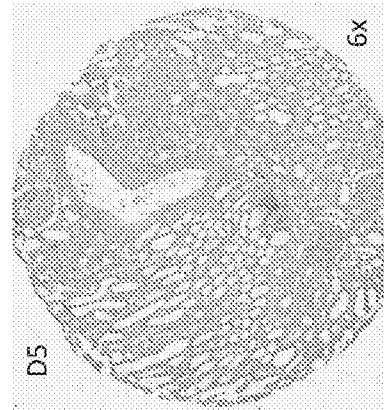


Fig. 194E

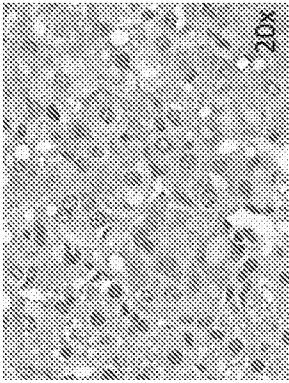


Fig. 194F



Fig. 194G

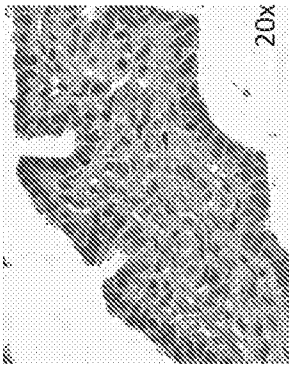


Fig. 194H



Figure 194A-194H

N+9/C-9 antibody 17H6 30 ug/mL  
FDA Normal tissue array MNO1021

Fig. 194I  
Normal Heart

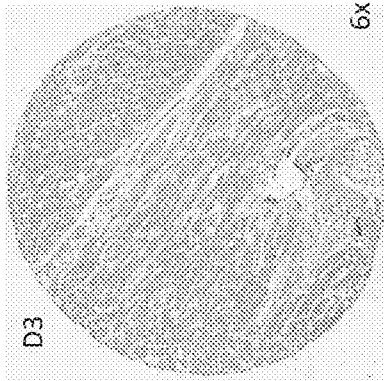


Fig. 194J  
Normal Liver

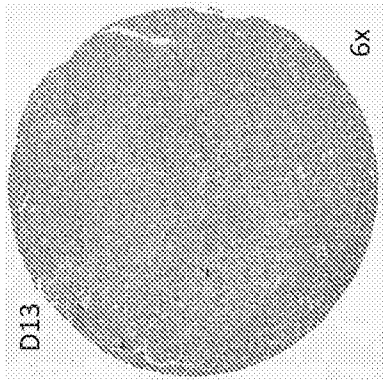


Fig. 194K  
Normal Lung

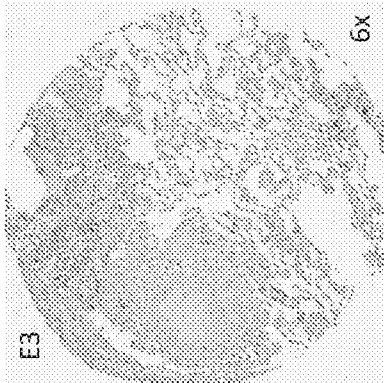


Fig. 194L  
Normal Ureter

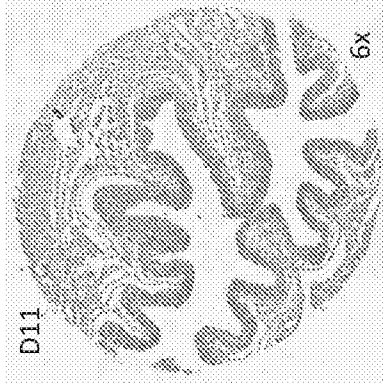


Fig. 194M

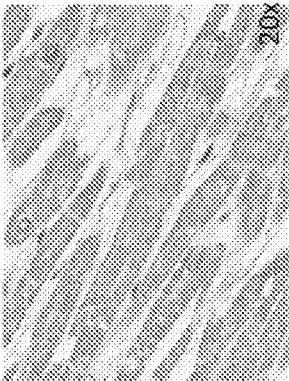


Fig. 194N

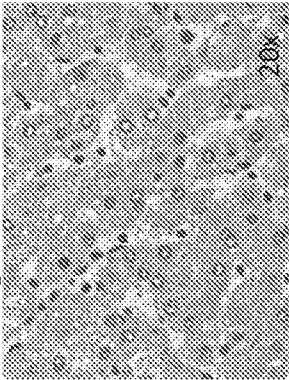


Fig. 194O

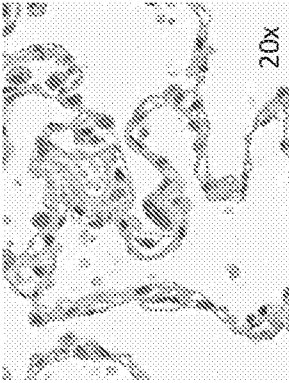


Fig. 194P

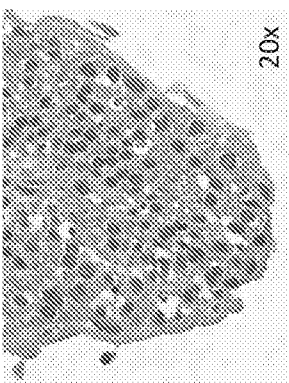


Figure 194I-194P

N+9/C-9 antibody 17H6 30 ug/mL  
FDA Normal tissue array MNO1021

Fig. 194Q  
Normal eye

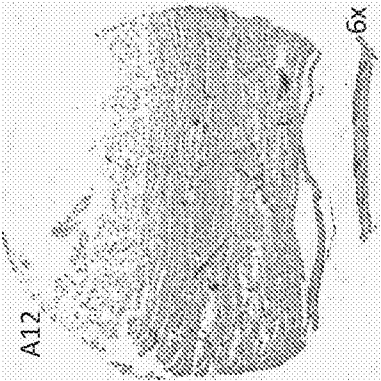


Fig. 194R  
Normal Cerebral cortex

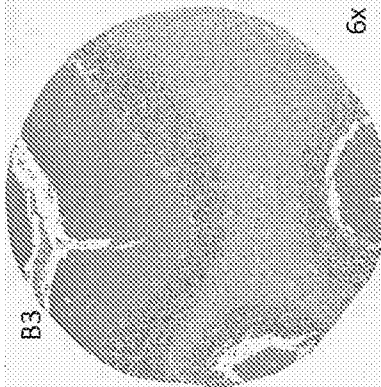


Fig. 194S  
Normal Bone marrow

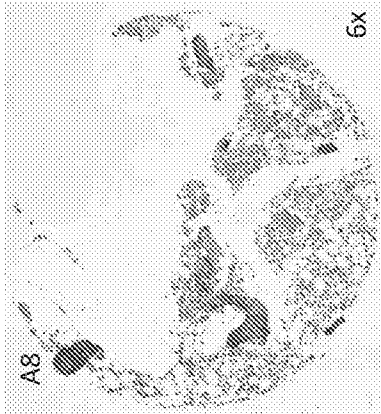


Fig. 194T  
Normal Skeletal muscle

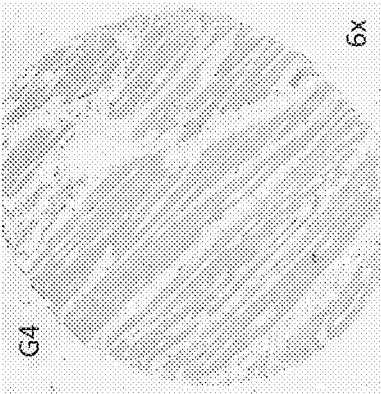


Fig. 194U

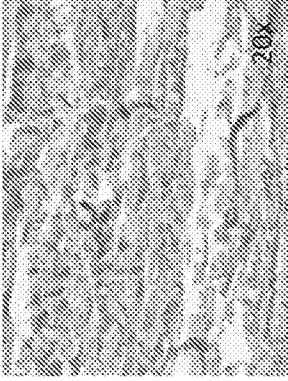


Fig. 194V

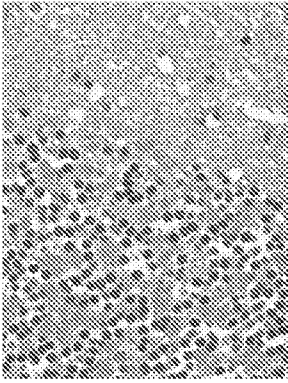


Fig. 194W

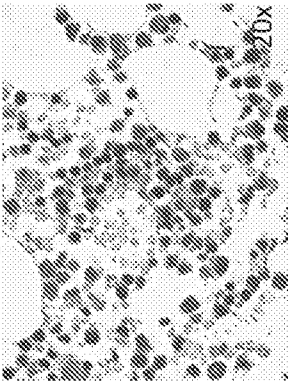


Fig. 194X

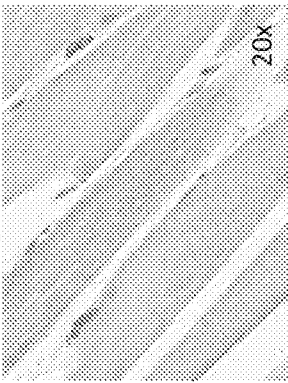


Figure 194Q-194X

351  
352  
353

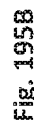
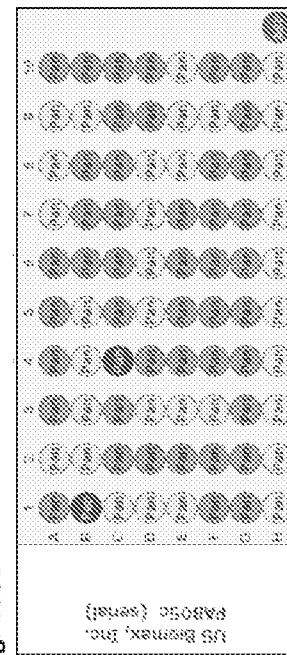



















Fig. 195A-195C

[illegible]

- Malignant tumor,  - Malignant tumor (stage 1)
- Malignant tumor,  - Malignant tumor (stage 2)
- Malignant tumor,  - Malignant tumor (stage 3)
- Malignant tumor,  - Malignant tumor (stage 4)
- Malignant tumor,  - Malignant tumor (stage 5)
- Malignant tumor,  - Malignant tumor (stage 6)
- Malignant tumor,  - Malignant tumor (stage 7)
- Malignant tumor,  - Malignant tumor (stage 8)
- Malignant tumor,  - Malignant tumor (stage 9)
- Malignant tumor,  - Malignant tumor (stage 10)
- Malignant tumor,  - Malignant tumor (stage 11)
- Malignant tumor,  - Malignant tumor (stage 12)
- Malignant tumor,  - Malignant tumor (stage 13)
- Malignant tumor,  - Malignant tumor (stage 14)
- Malignant tumor,  - Malignant tumor (stage 15)
- Malignant tumor,  - Malignant tumor (stage 16)
- Malignant tumor,  - Malignant tumor (stage 17)



N+9/C-9 antibody 17H6 IgG 30 ug/mL  
Pancreatic cancer tissue array PA805c

Fig. 196A

Position: D7  
Cell Type: Papillary Adenocarcinoma  
Tumor Grade: 2  
TNM: T2N0M0

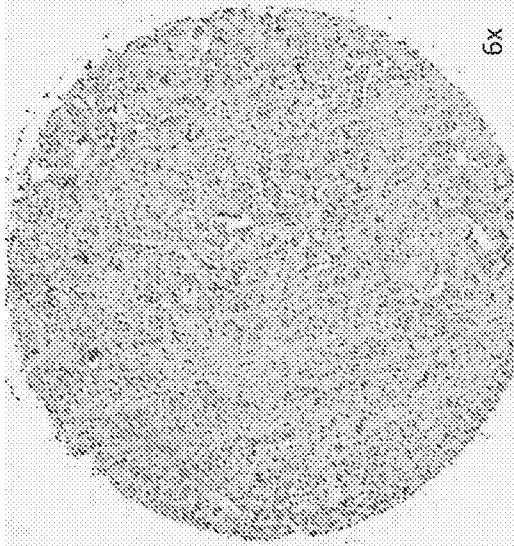


Fig. 196B

Position: E5  
Cell Type: Ductal Carcinoma  
Tumor Grade: 2-3  
TNM: T3N1M0

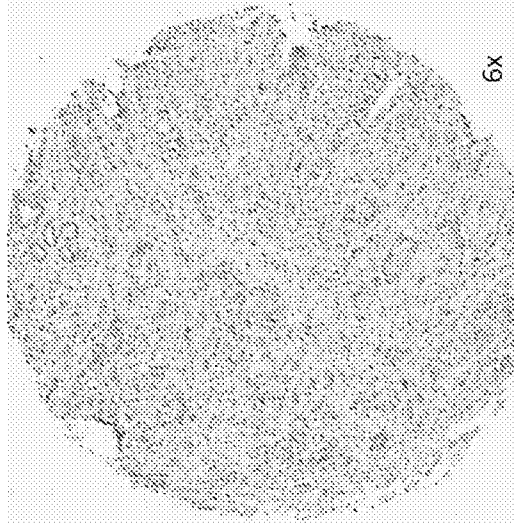


Fig. 196C

Position: F1  
Cell Type: Adenocarcinoma  
Tumor Grade: 3  
TNM: T3N0M0

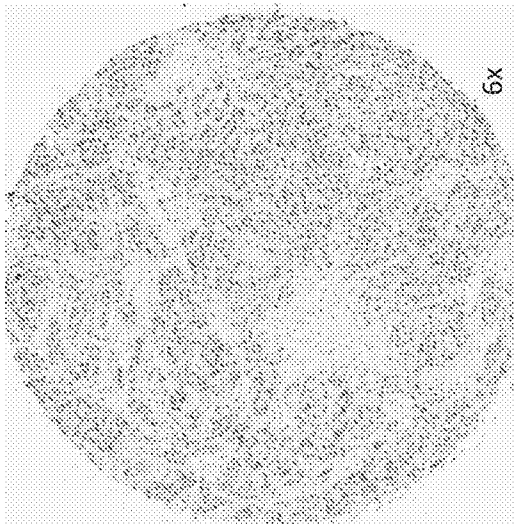


Fig. 196D



Fig. 196E

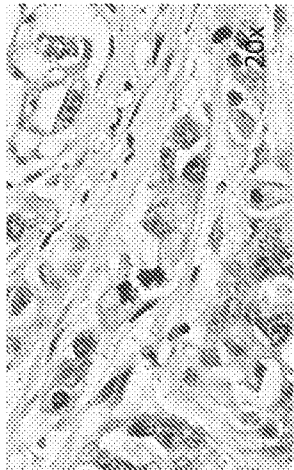


Fig. 196F

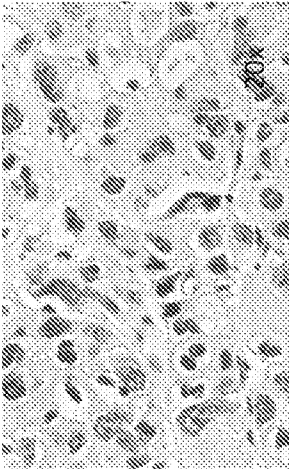


Fig. 196A-196F

N+9/C-9 antibody 39H5 5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 197A

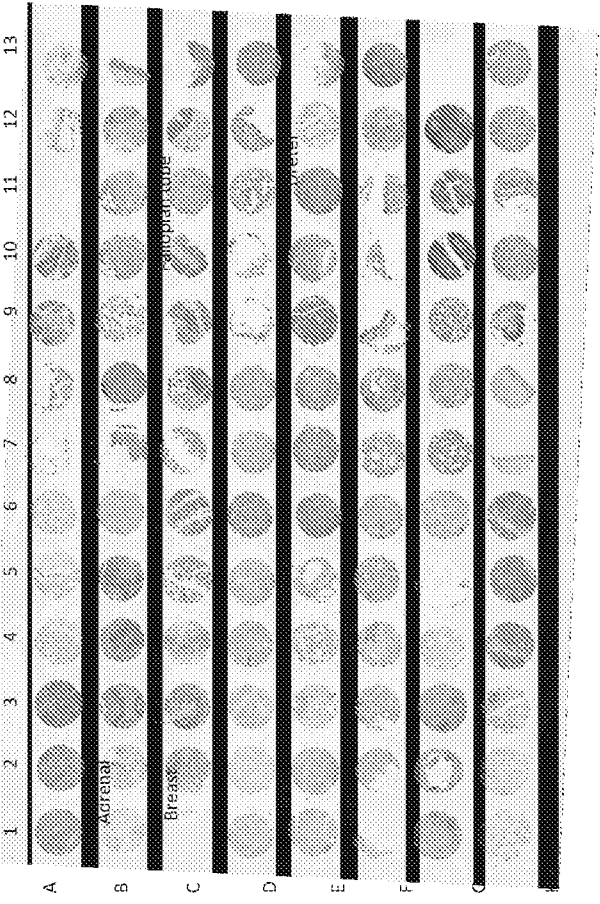


Fig. 197B

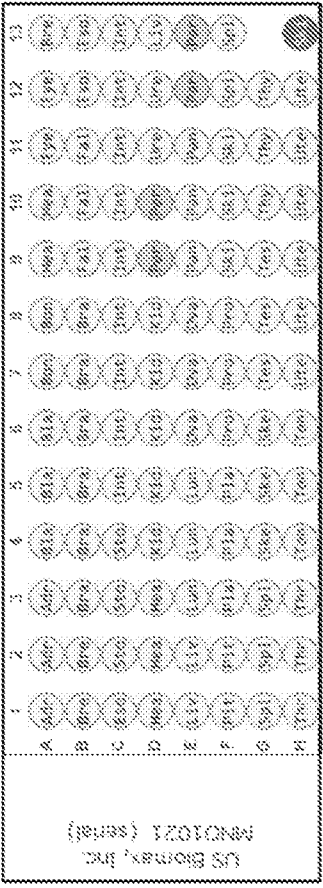


Figure 197A-197C

Fig. 197C

Position	Age	Sex	Organ/Anatomic Site	Pathology	Position	Age	Sex	Organ/Anatomic Site	Pathology
A1	F	31	Adrenal gland	Normal	E1	M	59	Liver	Normal
A2	M	51	Adrenal gland	Normal	E2	F	27	Liver	Normal
A3	M	53	Adrenal gland	Normal	E3	M	72	Liver	Normal
A4	M	73	Adrenal gland	Normal	E4	F	55	Lung	Normal
A5	M	73	Bladder, urinary	Normal	E5	M	43	Lung	Normal
A6	M	52	Bladder, urinary	Normal	E6	F	47	Ovary	Normal
A7	M	53	Bone, bone marrow	Normal	E7	F	46	Ovary	Normal
A8	F	44	Bone, bone marrow	Normal	E8	F	40	Ovary	Normal
A9	F	18	Head and neck, salivary gland	Normal	E9	M	42	Pancreas	Normal
A10	M	18	Head and neck, salivary gland	Normal	E10	F	35	Pancreas	Normal
A11	M	55	Eye	Normal	E11	F	59	Pancreas	Normal
A12	M	35	Eye	Normal	E12	M	17	Parathyroid	Adenoma
B1	F	35	Breast	Normal	E13	M	72	Parathyroid	Adenoma
B2	F	38	Breast	Normal	F1	F	3	Pituitary gland	Normal
B3	F	35	Breast	Normal	F2	F	32	Pituitary gland	Normal
B4	M	35	Brain, cerebellum	Normal	F3	F	30	Placenta	Normal
B5	M	35	Brain, cerebellum	Normal	F4	F	27	Placenta	Normal
B6	F	58	Brain, cerebral cortex	Normal	F5	M	64	Prostate	Normal
B7	M	85	Brain, cerebral cortex	Normal	F6	M	85	Prostate	Normal
B8	F	42	Fallopian tube	Normal	F7	M	32	Prostate	Normal
B9	F	32	Fallopian tube	Normal	F8	F	32	Prostate	Normal
B10	F	24	Fallopian tube	Normal	F9	F	38	Skin	Normal
B11	F	24	Fallopian tube	Normal	F10	M	25	Skin	Normal
B12	F	35	Esophagus	Normal	F11	M	47	Spinal cord	Normal
B13	F	35	Esophagus	Normal	F12	M	56	Spinal cord	Normal
C1	M	34	Esophagus	Normal	G1	M	27	Spleen	Normal
C2	M	49	Stomach	Normal	G2	M	90	Spleen	Normal
C3	M	49	Stomach	Normal	G3	M	31	Spleen	Normal
C4	M	77	Stomach	Normal	G4	F	49	Skeletal muscle	Normal
C5	F	73	Intestine, small intestine	Normal	G5	M	70	Skeletal muscle	Normal
C6	F	73	Intestine, small intestine	Normal	G6	M	43	Testis	Normal
C7	F	73	Intestine, small intestine	Normal	G7	M	30	Testis	Normal
C8	M	2	Intestine, small intestine	Normal	G8	M	30	Testis	Normal
C9	M	48	Intestine, colon	Normal	G9	M	15	Thyroid	Normal
C10	M	74	Intestine, colon	Normal	G10	M	28	Thyroid	Normal
C11	F	47	Intestine, rectum	Normal	G11	M	28	Thyroid	Normal
C12	F	75	Intestine, rectum	Normal	G12	F	9	Thyroid	Normal
C13	M	86	Intestine, rectum	Normal	G13	F	26	Thyroid	Normal
D1	M	38	Heart	Normal	H1	F	26	Thyroid	Normal
D2	M	38	Heart	Normal	H2	F	57	Thyroid	Normal
D3	F	34	Heart	Normal	H3	F	51	Thyroid	Normal
D4	F	32	Kidney, cortex	Normal	H4	M	46	Tonsil	Normal
D5	M	23	Kidney, cortex	Normal	H5	M	37	Tonsil	Normal
D6	M	52	Kidney, cortex	Normal	H6	M	38	Uterus, cervix	Normal
D7	M	52	Kidney, medulla	Normal	H7	F	35	Uterus, cervix	Normal
D8	M	23	Kidney, medulla	Schwannoma	H8	F	44	Uterus, cervix	Normal
D9	M	39	Peripheral nerve	Schwannoma	H9	F	38	Uterus, endometrium	Normal
D10	F	29	Peripheral nerve	Schwannoma	H10	F	38	Uterus, endometrium	Normal
D11	F	44	Ureter	Normal	H11	F	41	Uterus, endometrium	Normal
D12	F	42	Ureter	Normal	H12	F	46	Uterus, endometrium	Normal
D13	F	57	Liver	Normal	H13	M	58	Skin	Malignant melanoma (tissue matrix)



N+9/C-9 antibody 39H5 5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 198A

Normal Adrenal Gland

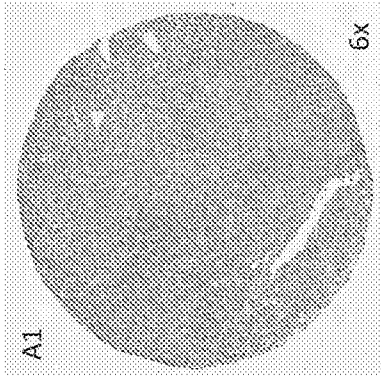


Fig. 198B

Normal Breast

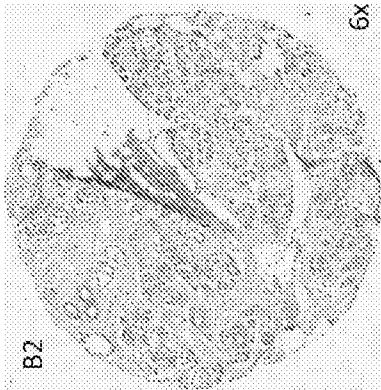


Fig. 198C

Normal Fallopian Tubes

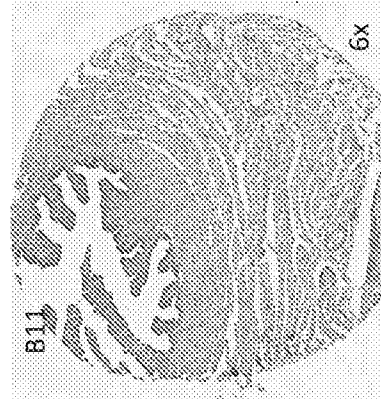


Fig. 198D

Normal Kidney

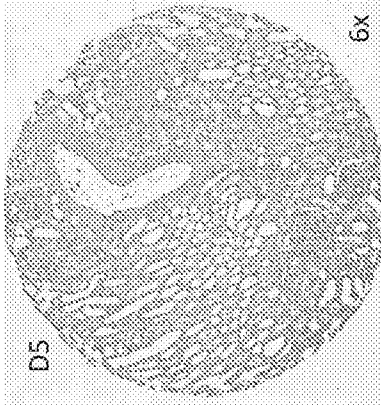


Fig. 198E

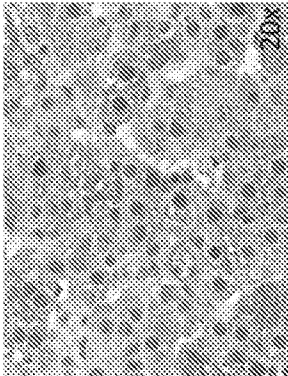


Fig. 198F



Fig. 198G

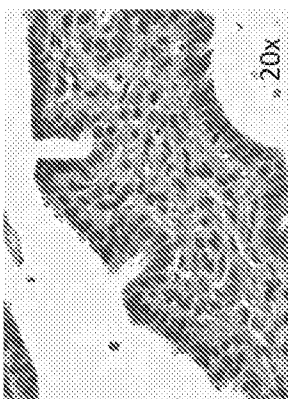


Fig. 198H



Figure 198A-198H

N+9/C-9 antibody 39H5 5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 198I  
Normal Heart

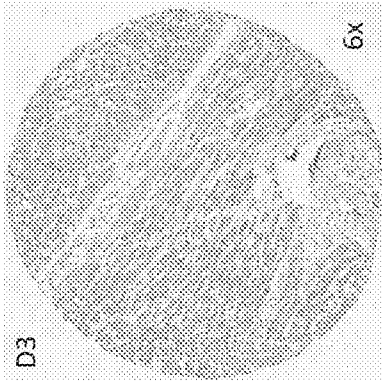


Fig. 198J  
Normal Liver

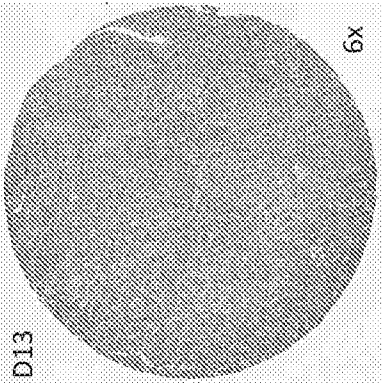


Fig. 198K  
Normal Lung

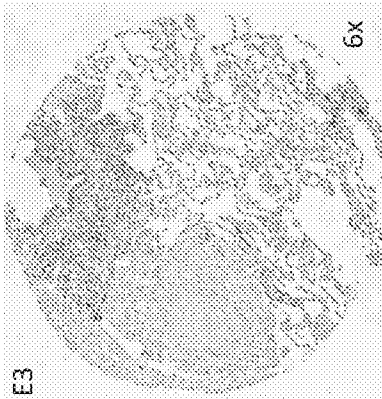


Fig. 198L  
Normal Ureter

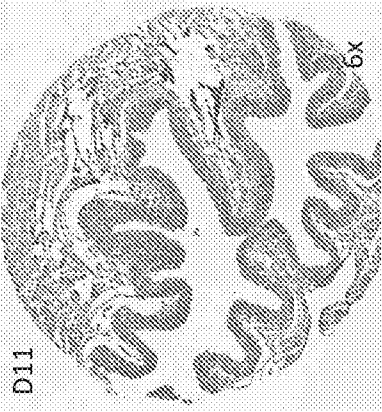


Fig. 198M

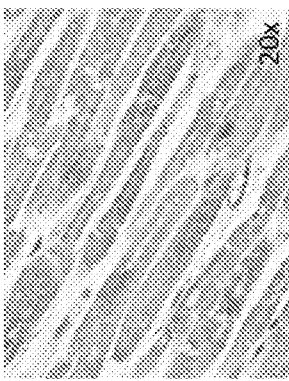


Fig. 198N

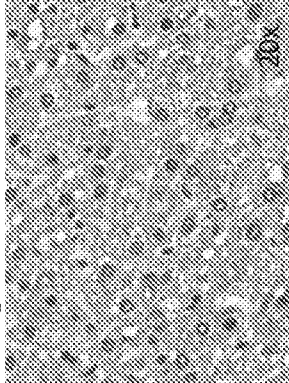


Fig. 198O

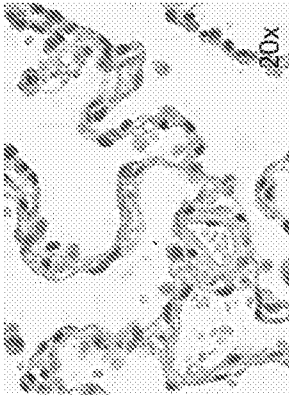


Fig. 198P



Figure 198I-198P

N+9/C-9 antibody 39H5 5 ug/mL  
FDA Normal tissue array MNO1021

Fig. 198Q  
Normal eye

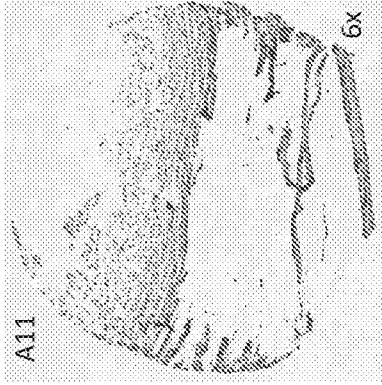


Fig. 198R  
Normal Cerebral cortex

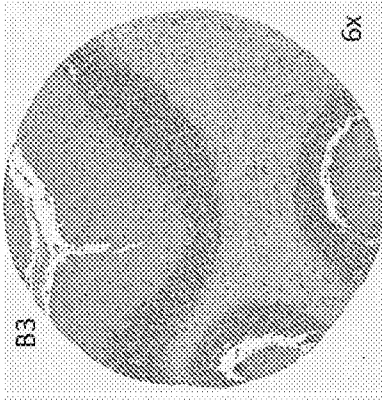


Fig. 198S  
Normal Bone marrow

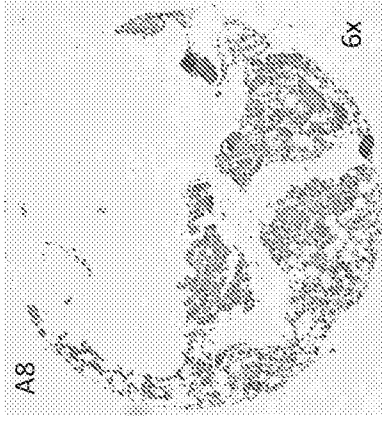


Fig. 198T  
Normal Skeletal muscle

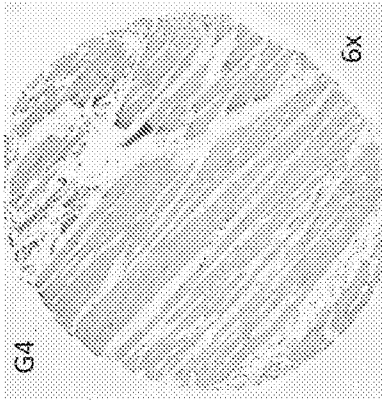


Fig. 198U

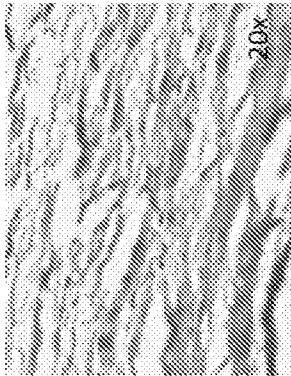


Fig. 198V

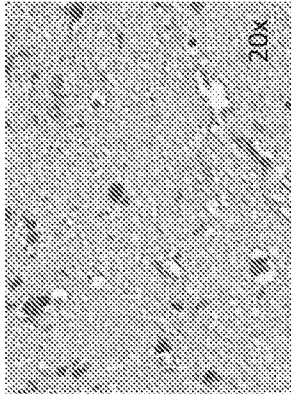


Fig. 198W

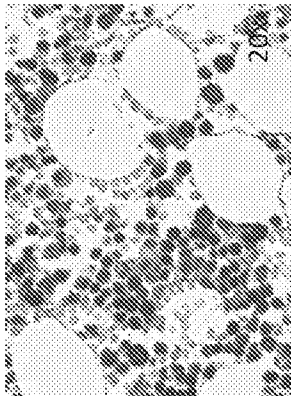


Fig. 198X

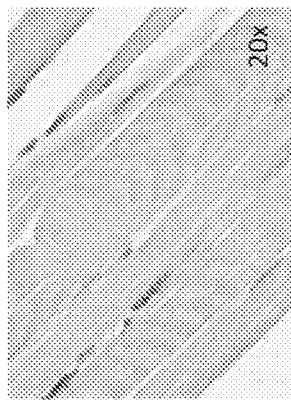


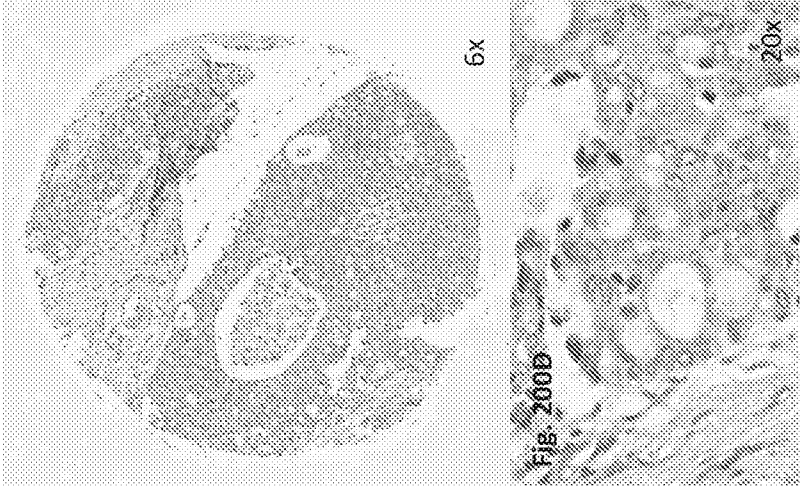
Figure 198Q-198X



**N+9/C-9 antibody 39H5 5 ug/mL**  
**Pancreatic cancer tissue array PA1003**

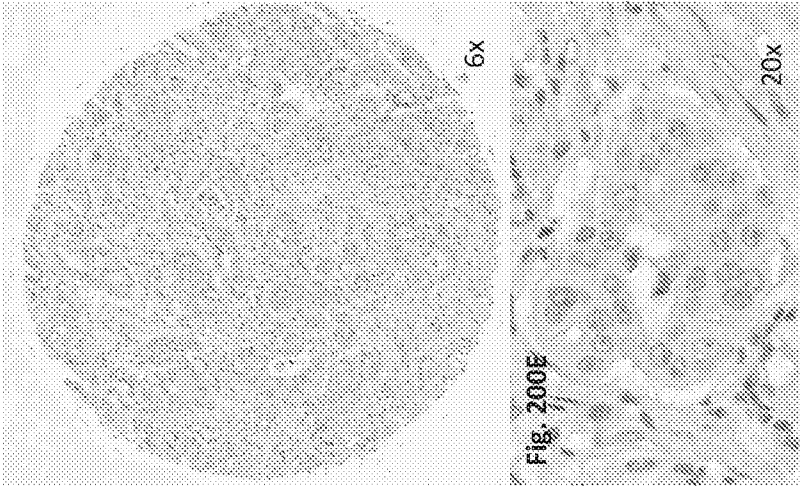
**Fig. 200A**

Position: F6  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



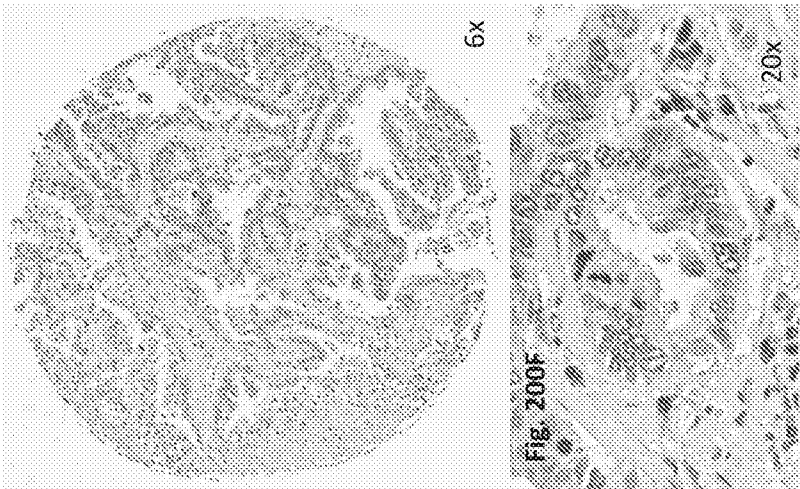
**Fig. 200B**

Position: F7  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



**Fig. 200C**

Position: G8  
Cell Type: Adenocarcinoma  
Tumor Grade: 2  
TNM: T3N0M0



**Fig. 200A-200F**

ELISA – binding of antibodies raised against PSMGFR

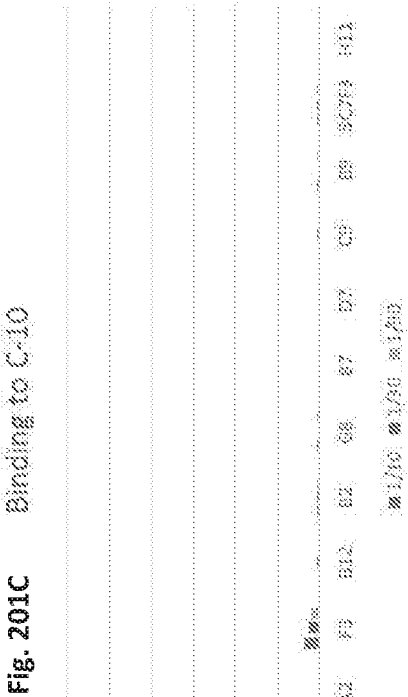
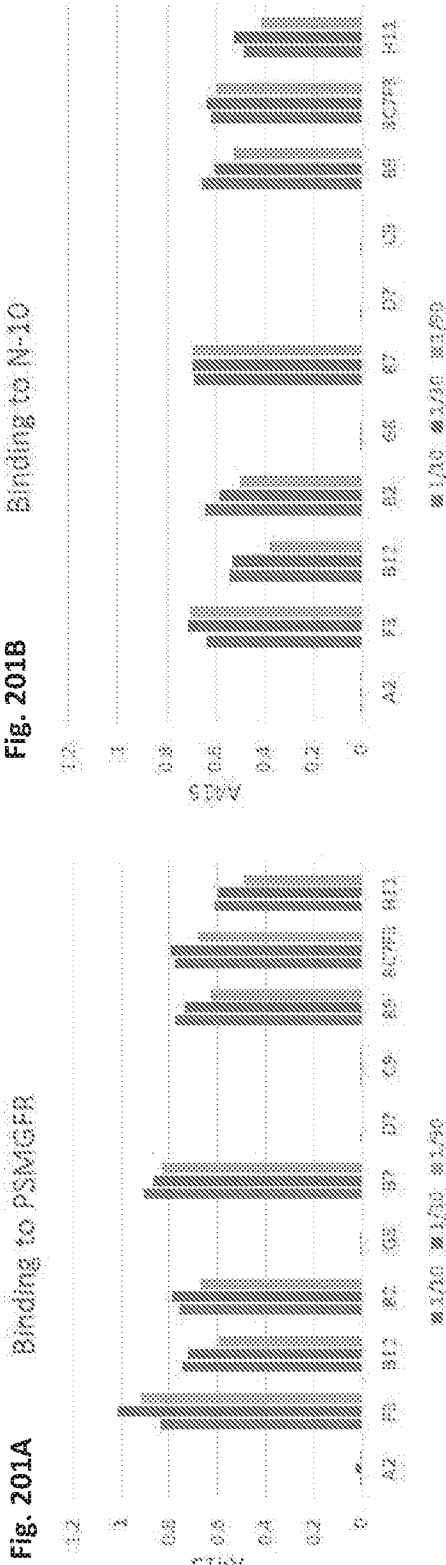


Figure 201A-201C

Pancreatic Array PA1003:

Fig. 202A

10 ug/mL 1E4 (N+20/C-27)

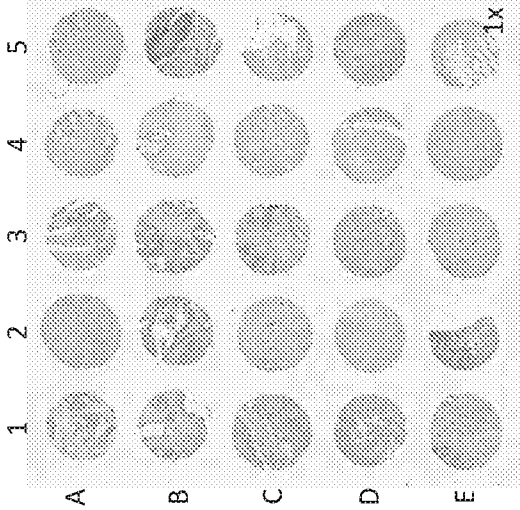


Fig. 202B

10 ug/mL 18B4 (PSMGFR)

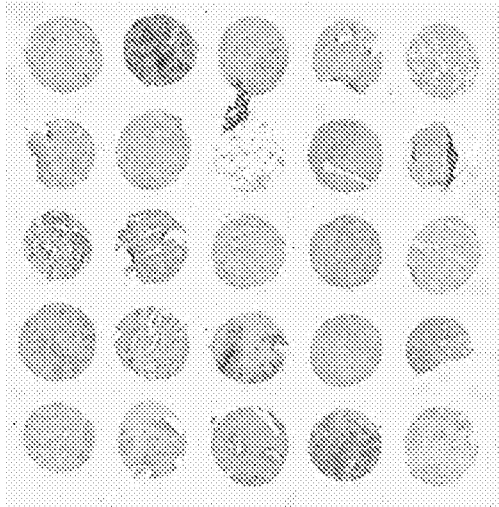


Fig. 202C

1 ug/mL SDIX (PSMGFR)

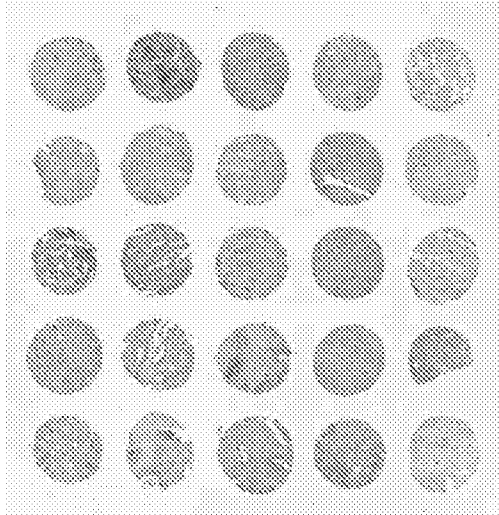


Figure 202A-202C



Pancreatic Array PA1003: Donor A2

Fig. 203A

10 ug/mL 1E4 (N+20/C-27)

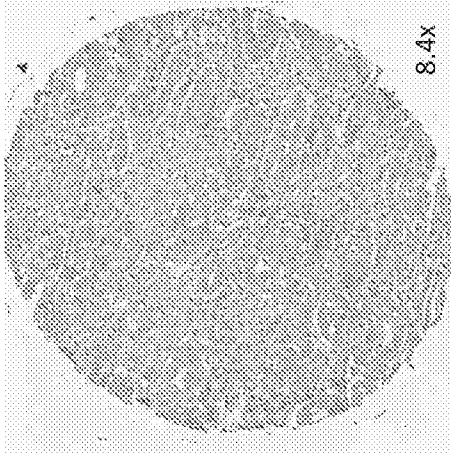


Fig. 203C

10 ug/mL 18B4 (PSMGFR)

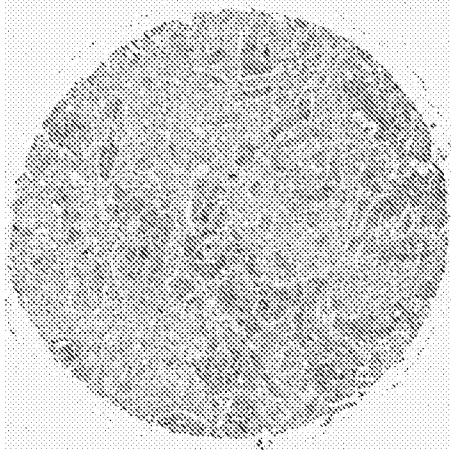


Fig. 203E

1 ug/mL SDIX (PSMGFR)

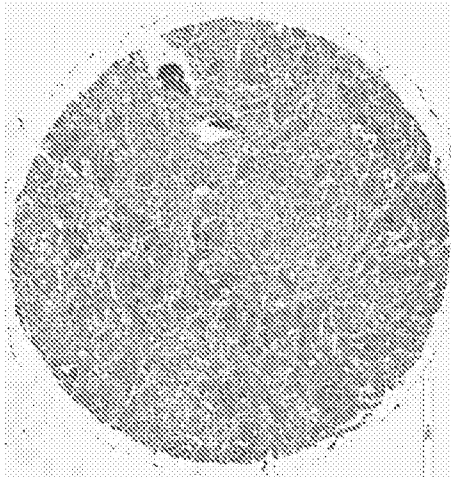


Fig. 203B

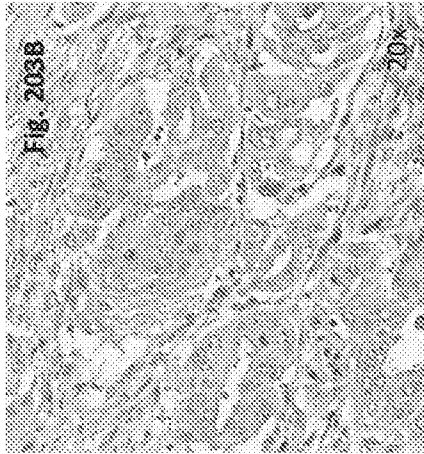


Fig. 203D

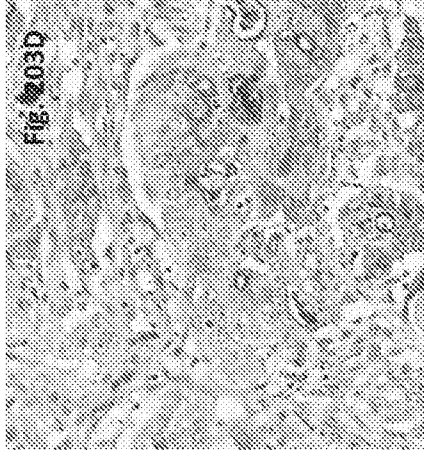


Fig. 203F

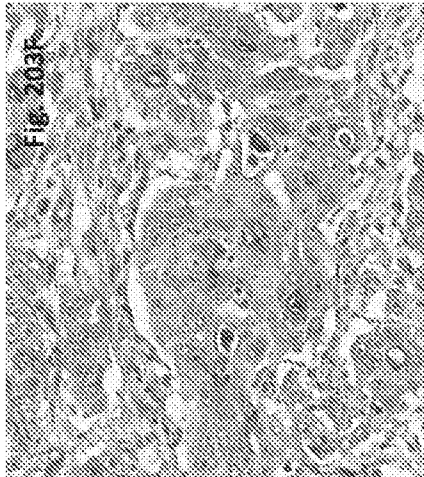


Figure 203A-203F



Pancreatic Array PA1003: Donor D4

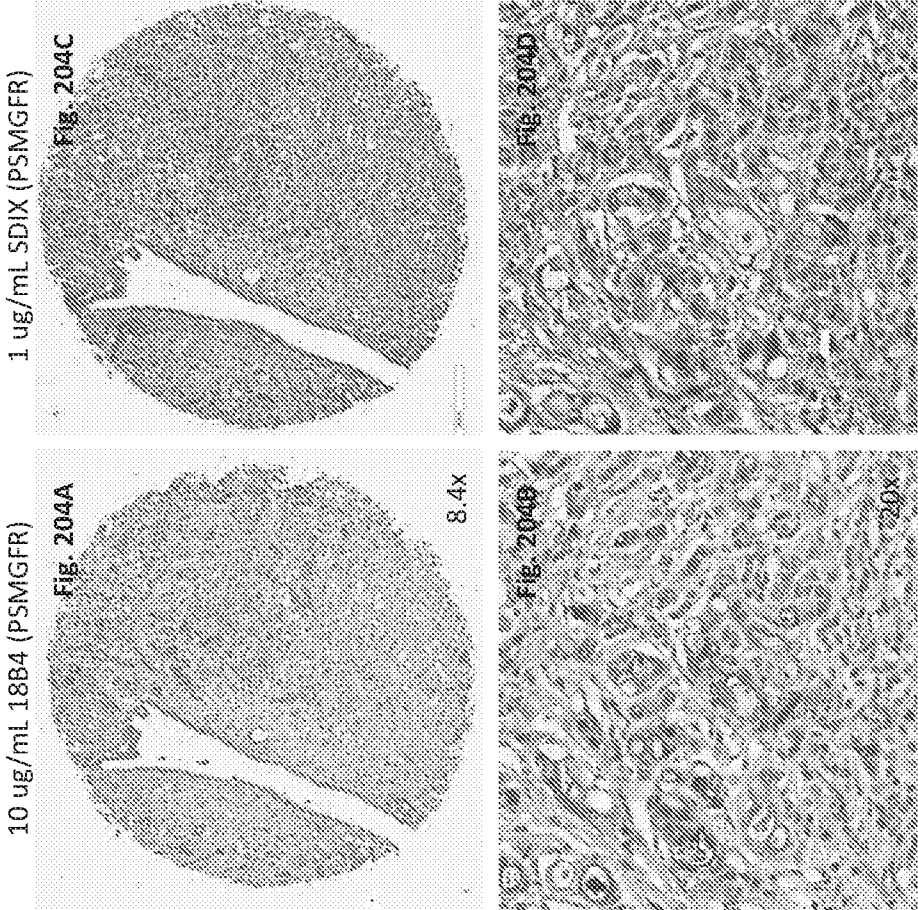


Figure 204A-204D

Pancreatic Array PA1003: Donor E1

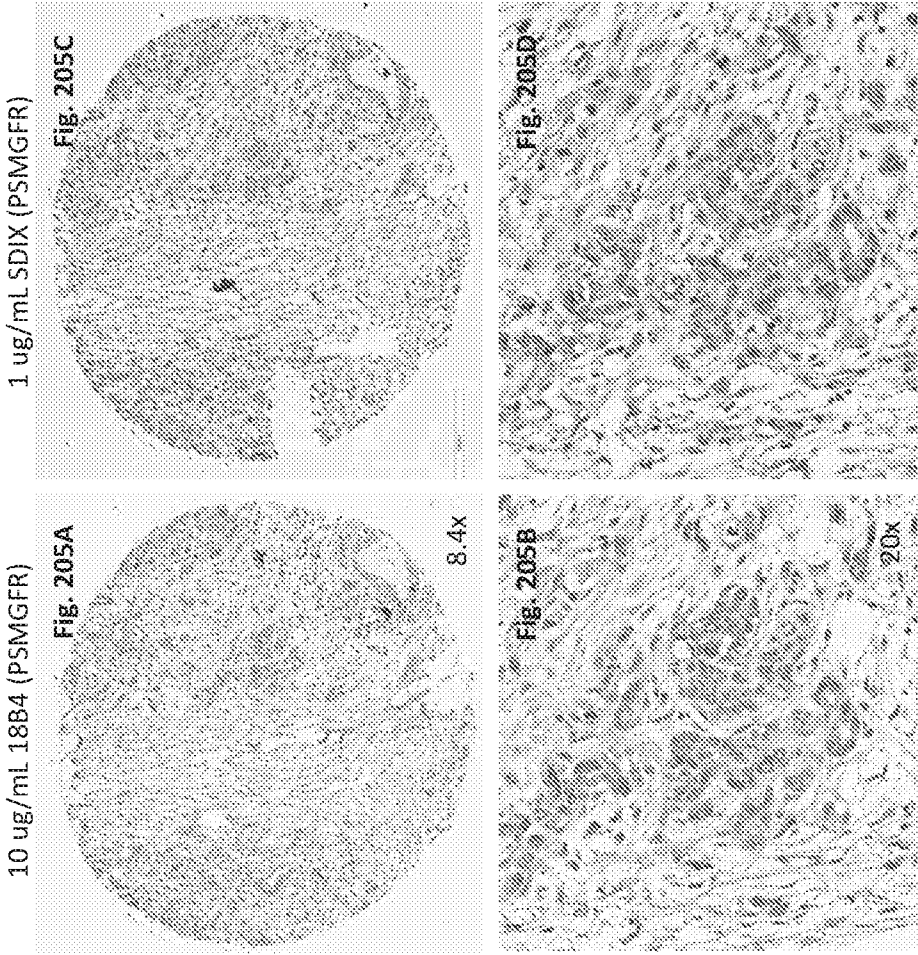


Figure 205A-205D

Pancreatic Array PA1003: Donor C3

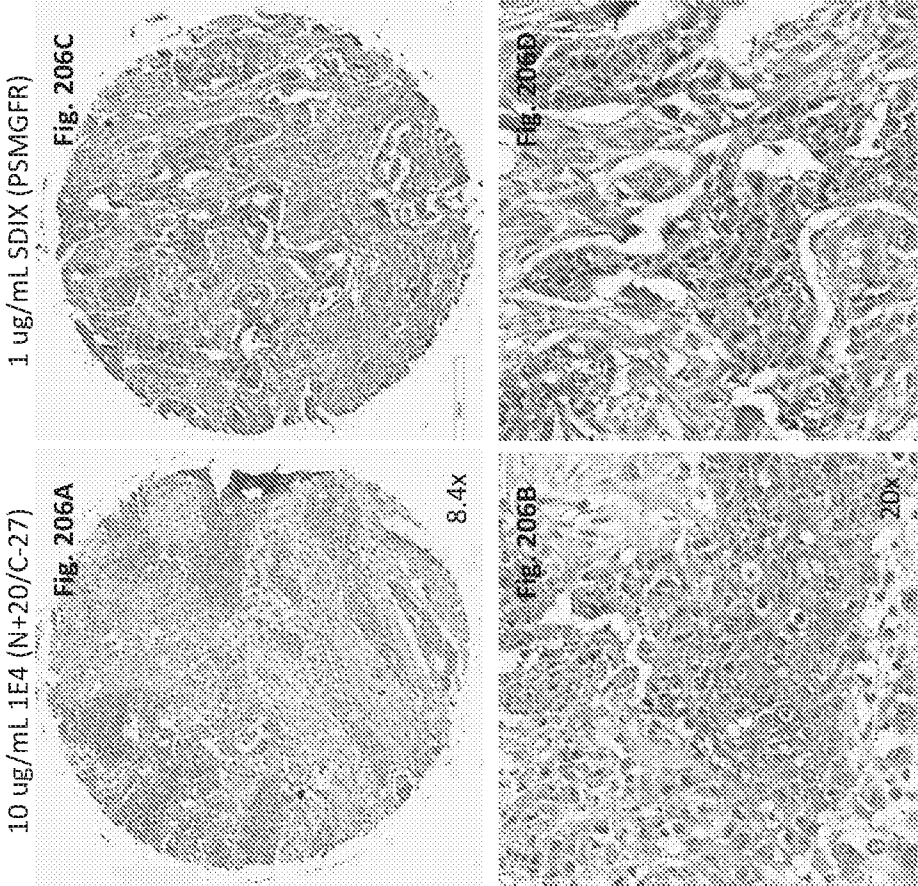


Figure 206A-206D

Pancreatic Array PA1003: Donor D1

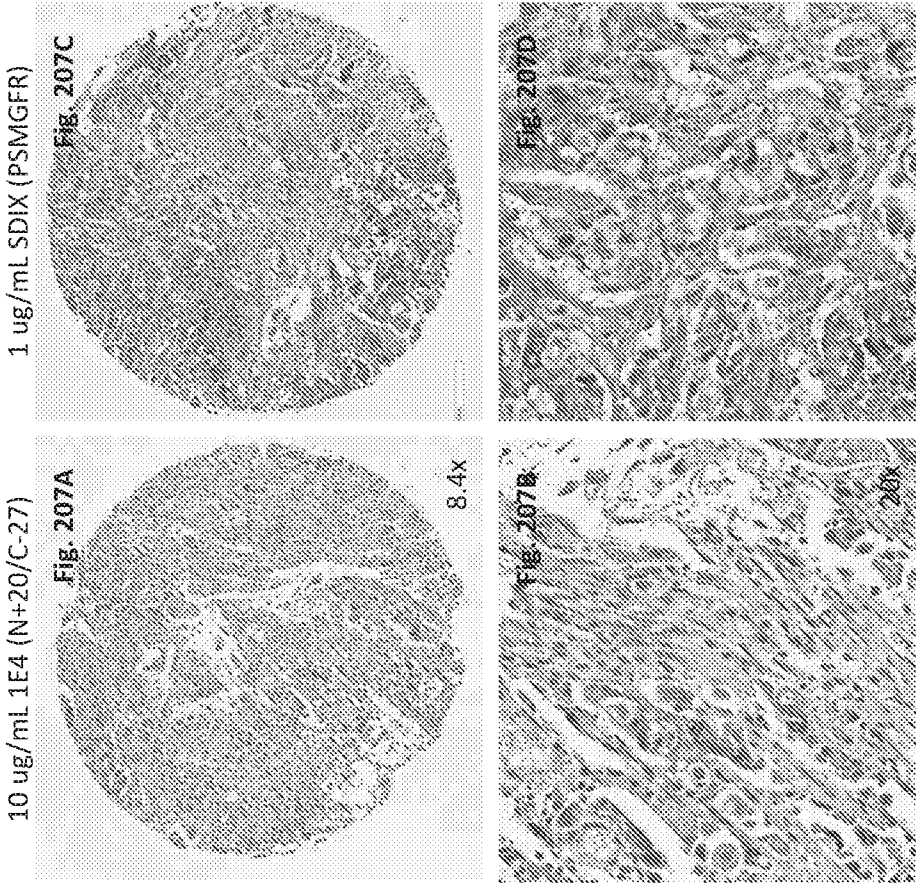
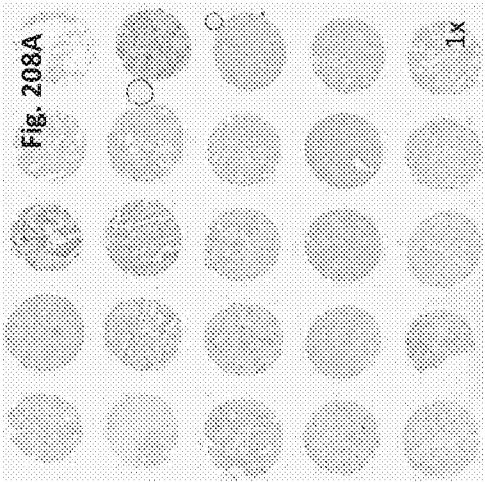


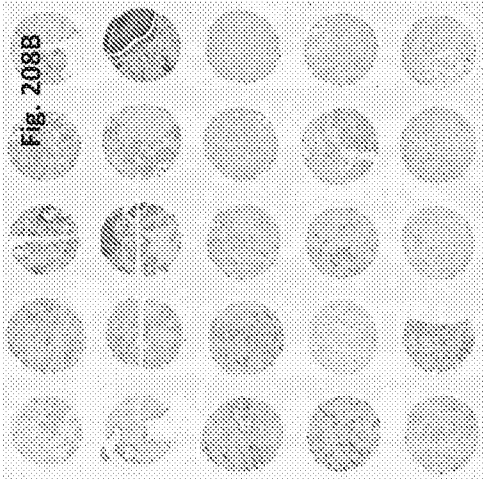
Figure 207A-207D

Pancreatic Array PA1003:

1 ug/mL SDIX (PSMGFR)



0.5 ug/mL 20A10 (PSMGFR)



4 ug/mL 29H1 (N+20/C-27)

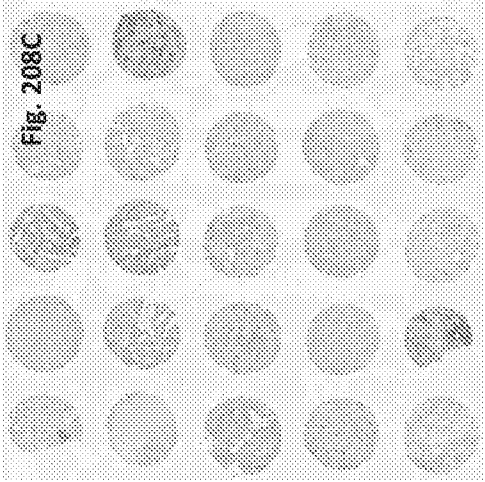
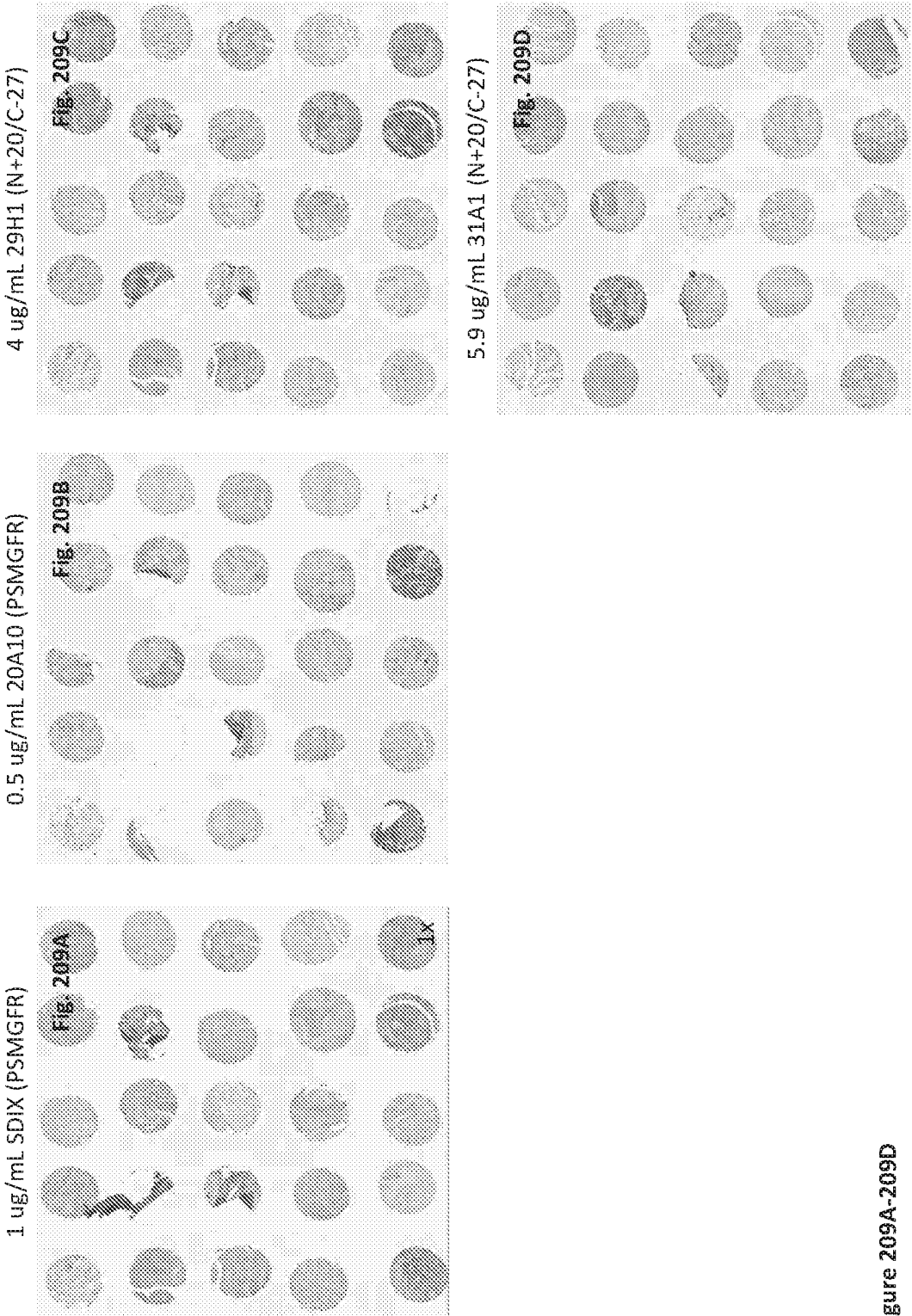


Figure 208A-208C

Esophageal Cancer Array ES1001: overview of MUC1\* monoclonal staining



Prostate Cancer Array PR1001: overview of MUC1\* monoclonal staining

1 ug/mL SDIX (PSMGFR)      0.5 ug/mL 20A10 (PSMGFR)      4 ug/mL 29H1 (N+20/C-27)

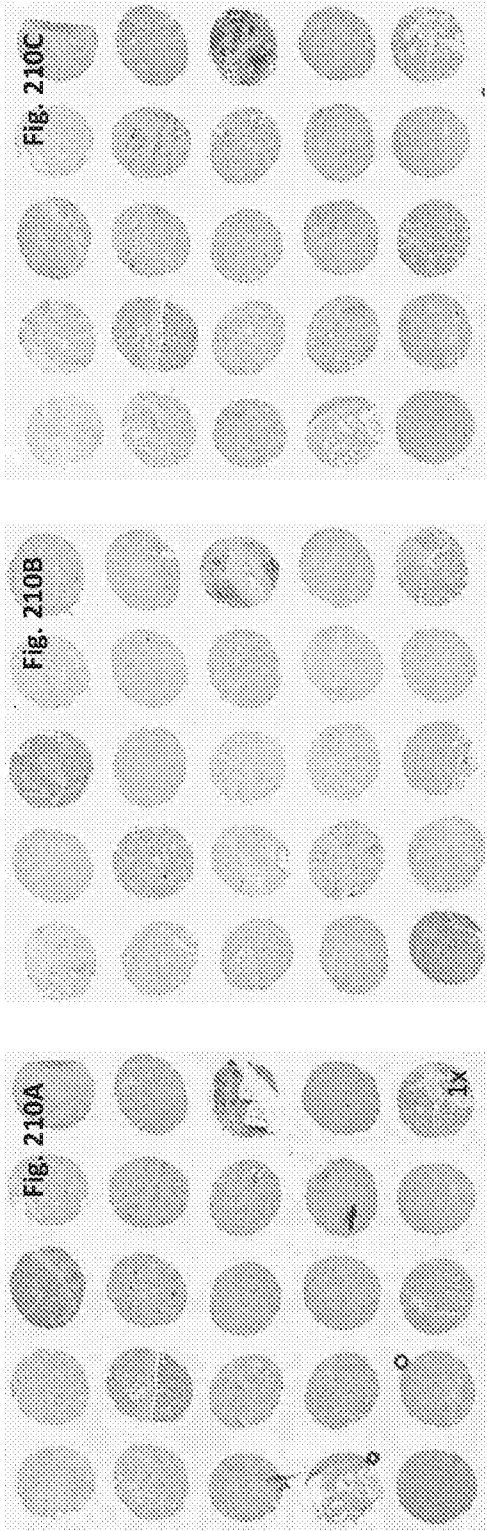


Figure 210A-210C

huMNC2-CAR44 with NFAT inducible IL-18 for increased persistence

Fig. 211A

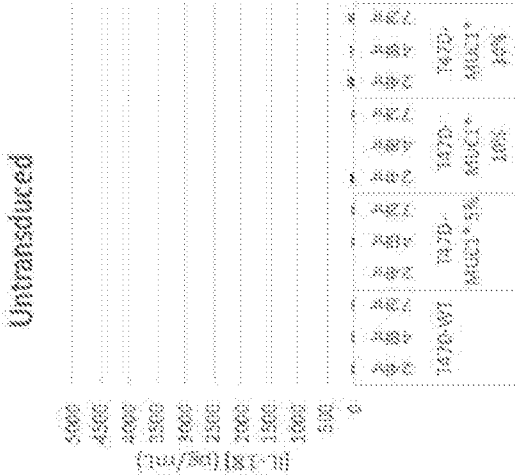


Fig. 211B

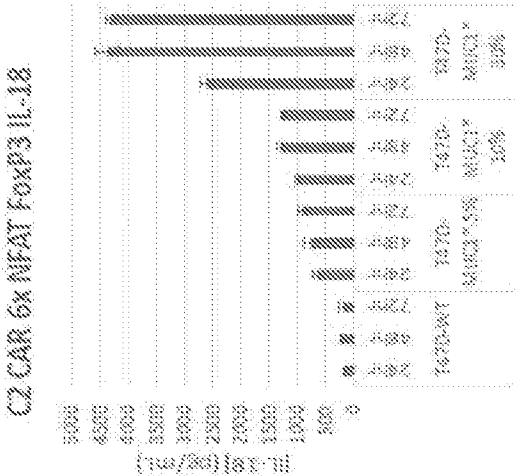
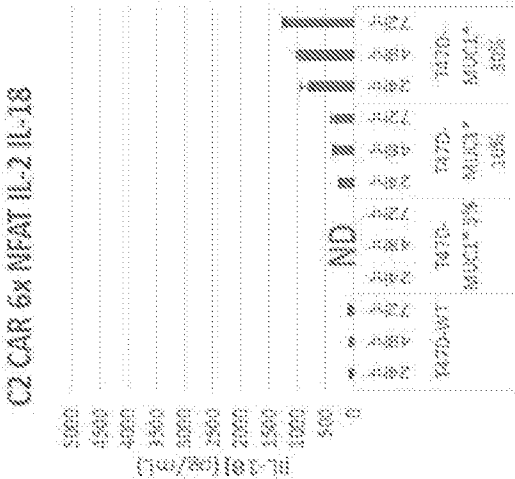


Fig. 211C



T47D breast cancer cells low – med antigen density; doped with increasing percentages of high antigen density cells

Figure 211A-211C



IL-18 also increases killing of low antigen density cells in a concentration dependent manner

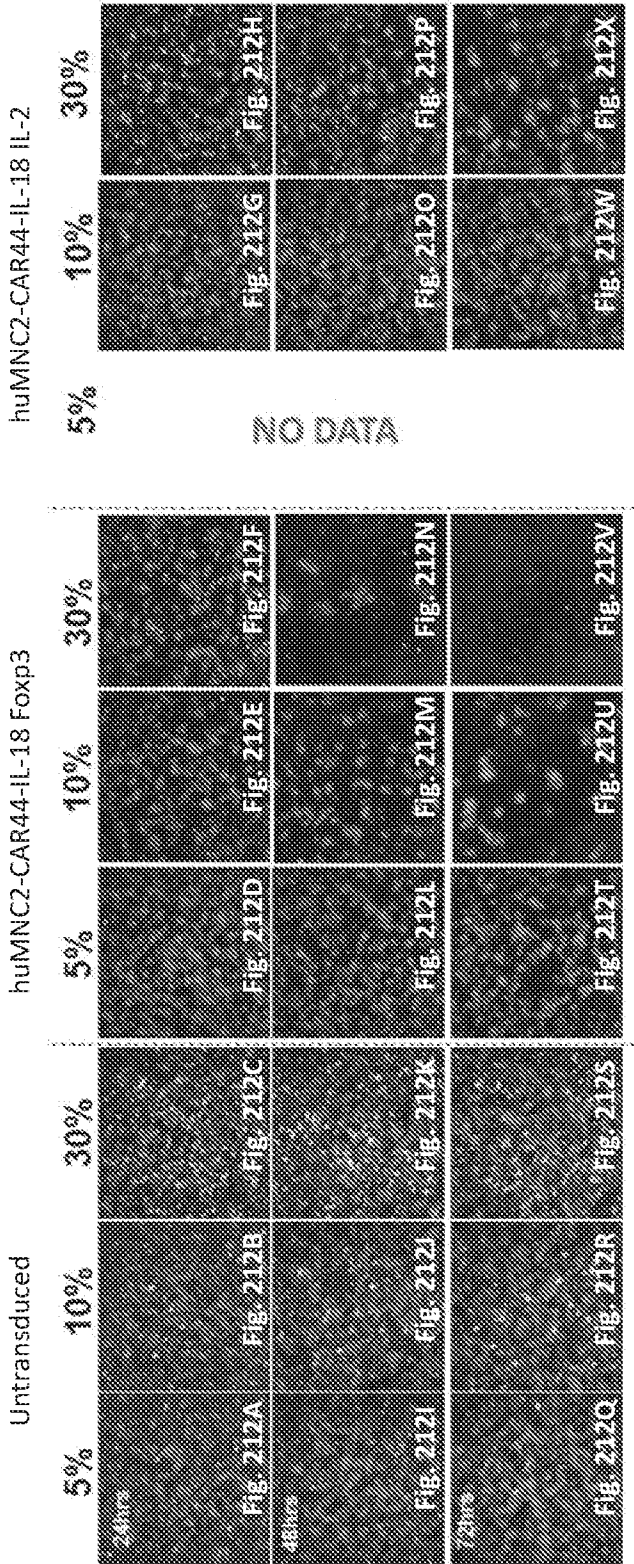


Figure 212A-212X

huMNC2-CAR44-IL-18 does not increase killing of normal cells

Fig. 213A

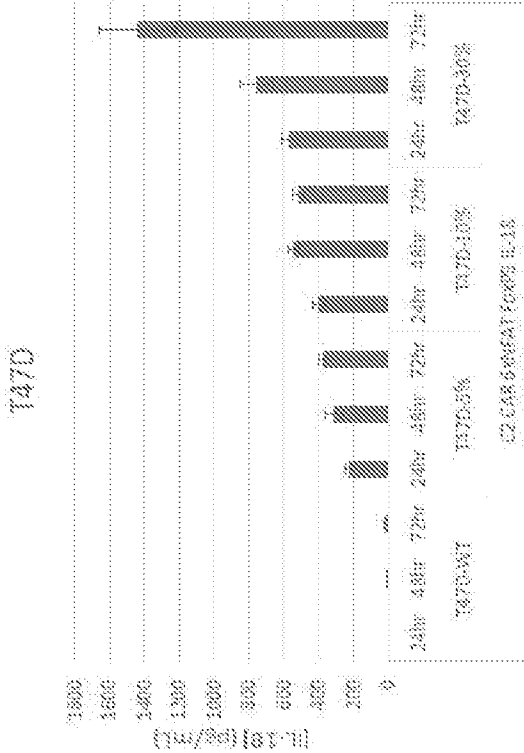


Fig. 213B

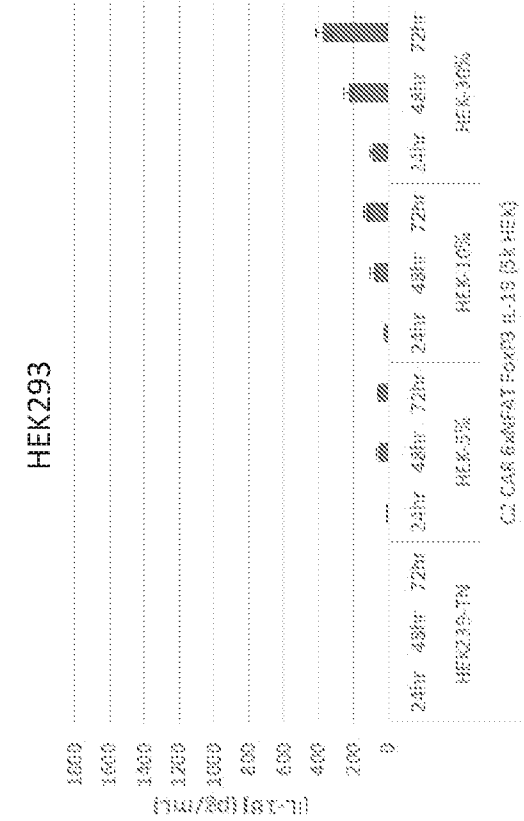


Figure 213A-213B

At 72 hrs still no killing of non-target cells, but all high density cells gone

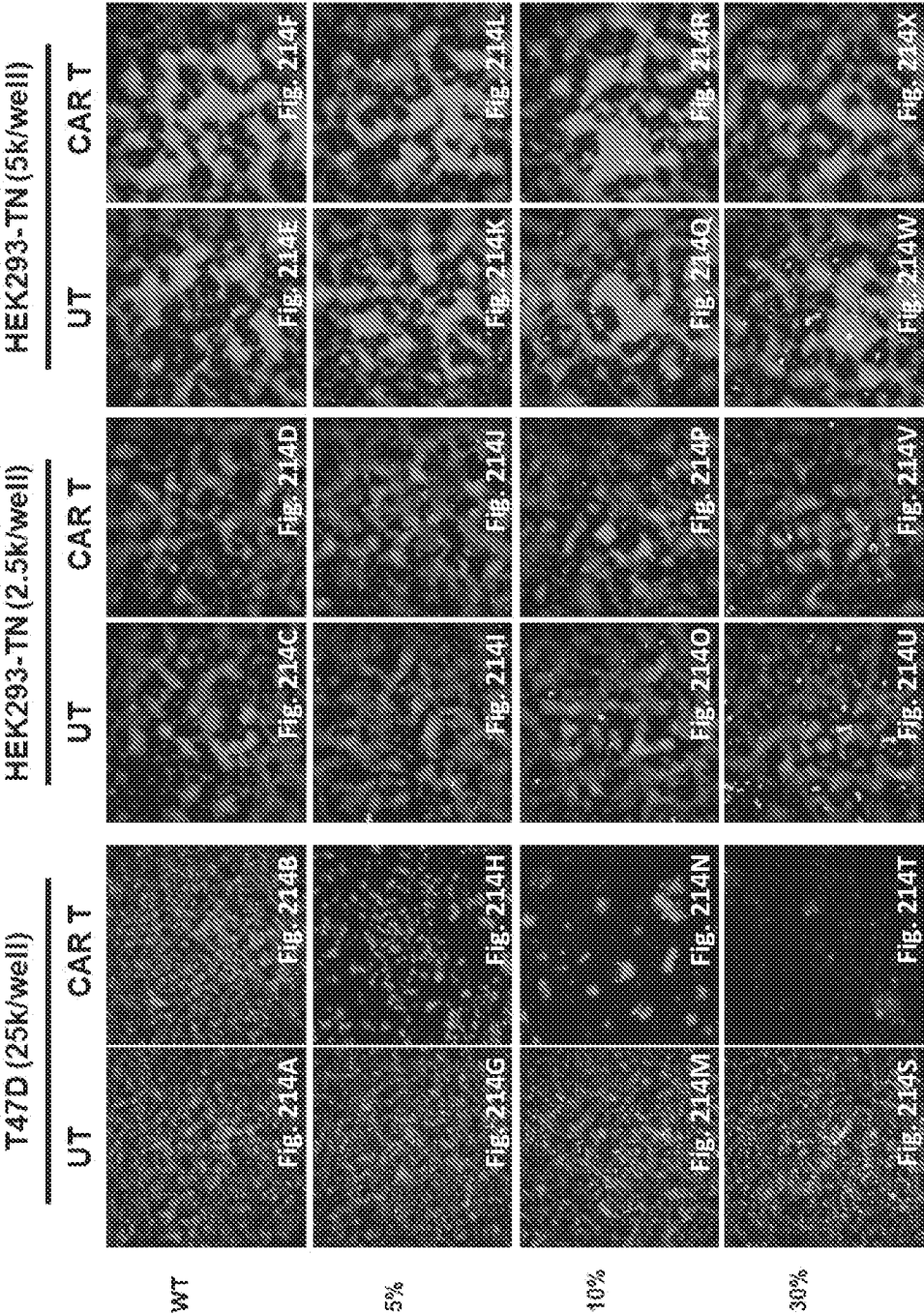
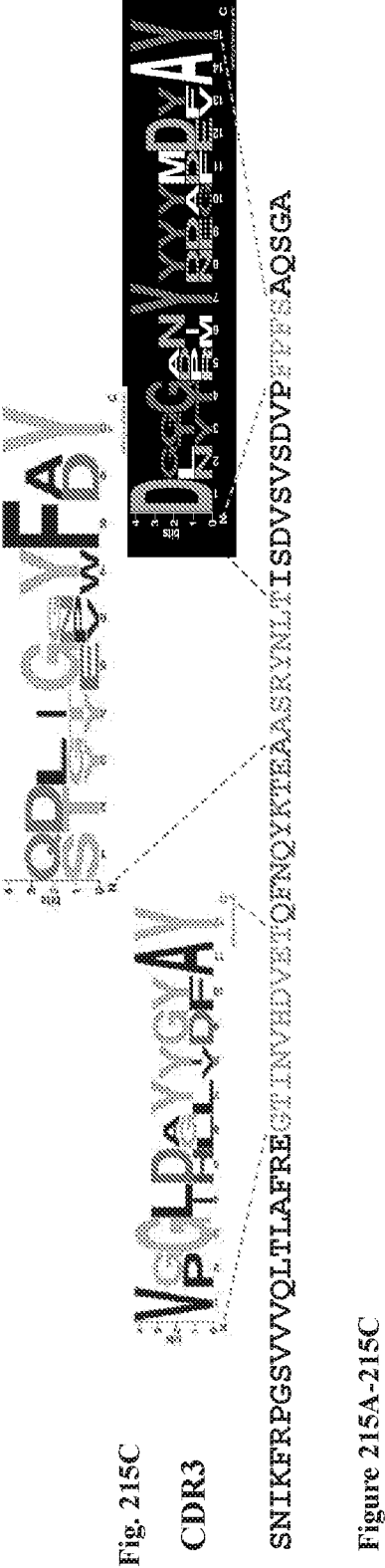
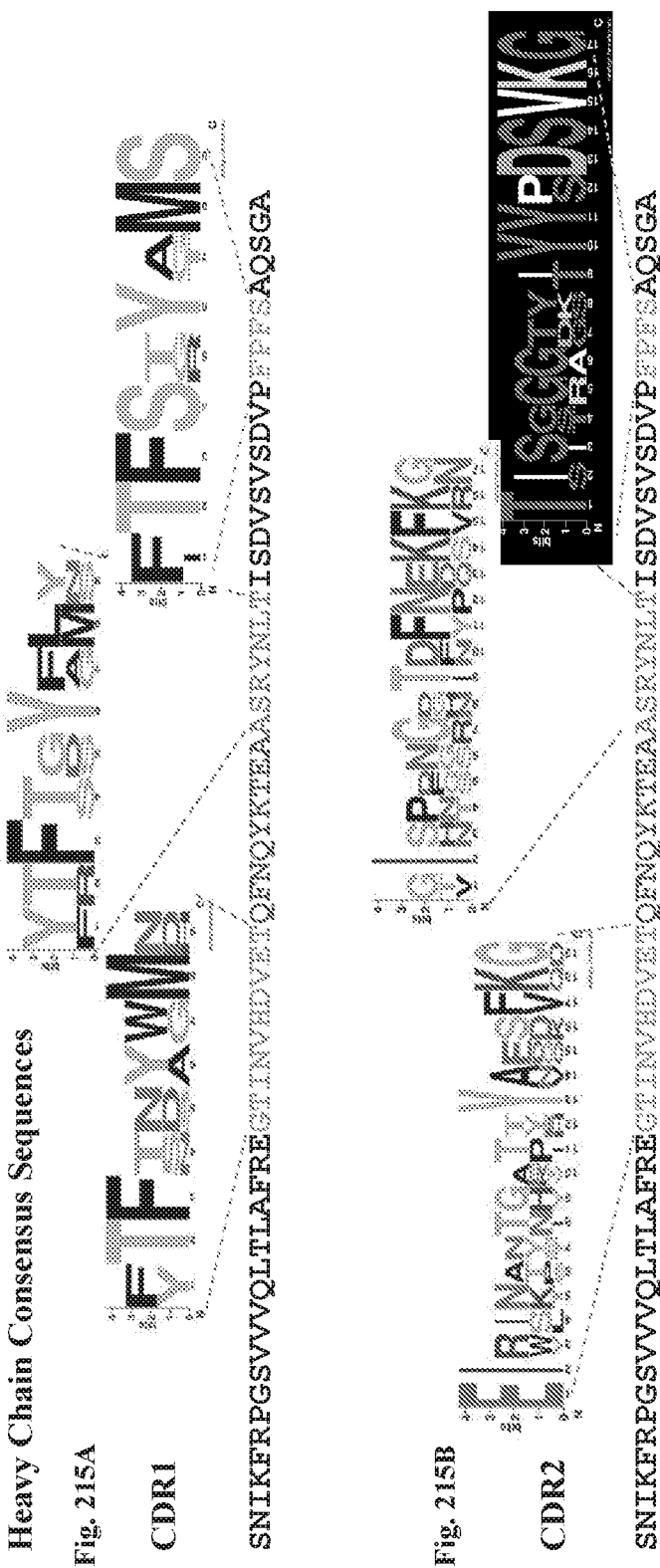


Figure 214A-214X



Light Chain Consensus Sequences

Fig. 216A



Fig. 216B

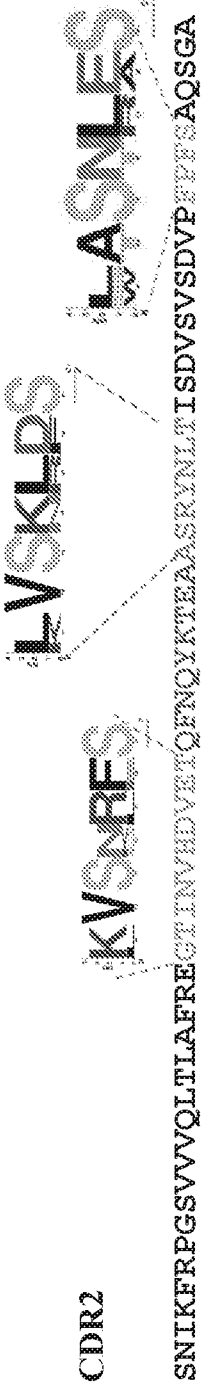


Fig. 216C



Figure 216A-216C

# SEQUENCE LISTING

<110> MINERVA BIOTECHNOLOGIES CORPORATION  
 <120> ANTI-VARIABLE MUC1\* ANTIBODIES AND USES THEREOF  
 <130> 6977-5801  
 <140> PCT/US2020/013410  
 <141> 2020-01-13  
 <160> 1790  
 <170> PatentIn version 3.5  
 <210> 1  
 <211> 1255  
 <212> PRT  
 <213> Homo sapiens  
 <400> 1

Met Thr Pro Gly Thr Gln Ser Pro Phe Phe Leu Leu Leu Leu Thr  
 1 5 10 15

Val Leu Thr Val Val Thr Gly Ser Gly His Ala Ser Ser Thr Pro Gly  
 20 25 30

Gly Glu Lys Glu Thr Ser Ala Thr Gln Arg Ser Ser Val Pro Ser Ser  
 35 40 45

Thr Glu Lys Asn Ala Val Ser Met Thr Ser Ser Val Leu Ser Ser His  
 50 55 60

Ser Pro Gly Ser Gly Ser Ser Thr Thr Gln Gly Gln Asp Val Thr Leu  
 65 70 75 80

Ala Pro Ala Thr Glu Pro Ala Ser Gly Ser Ala Ala Thr Trp Gly Gln  
 85 90 95

Asp Val Thr Ser Val Pro Val Thr Arg Pro Ala Leu Gly Ser Thr Thr  
 100 105 110

Pro Pro Ala His Asp Val Thr Ser Ala Pro Asp Asn Lys Pro Ala Pro  
 115 120 125

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
 130 135 140

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
 145 150 155 160

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
 165 170 175

Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
 180 185 190

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
195 200 205

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
210 215 220

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
225 230 235 240

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
245 250 255

Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
260 265 270

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
275 280 285

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
290 295 300

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
305 310 315 320

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
325 330 335

Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
340 345 350

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
355 360 365

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
370 375 380

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
385 390 395 400

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
405 410 415

Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
420 425 430

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
435 440 445

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
450 455 460

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
465 470 475 480

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
485 490 495

Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
500 505 510

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
515 520 525

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
530 535 540

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
545 550 555 560

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
565 570 575

Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
580 585 590

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
595 600 605

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
610 615 620

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
625 630 635 640

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
645 650 655

Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
660 665 670

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
675 680 685

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
690 695 700

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
705 710 715 720

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
725 730 735



Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
740 745 750

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
755 760 765

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
770 775 780

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
785 790 795 800

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
805 810 815

Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
820 825 830

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
835 840 845

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr  
850 855 860

Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser  
865 870 875 880

Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His  
885 890 895

Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala  
900 905 910

Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro  
915 920 925

Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Asn  
930 935 940

Arg Pro Ala Leu Gly Ser Thr Ala Pro Pro Val His Asn Val Thr Ser  
945 950 955 960

Ala Ser Gly Ser Ala Ser Gly Ser Ala Ser Thr Leu Val His Asn Gly  
965 970 975

Thr Ser Ala Arg Ala Thr Thr Thr Pro Ala Ser Lys Ser Thr Pro Phe  
980 985 990

Ser Ile Pro Ser His His Ser Asp Thr Pro Thr Thr Leu Ala Ser His  
995 1000 1005

Ser Thr Lys Thr Asp Ala Ser Ser Thr His His Ser Ser Val Pro

1010		1015		1020
Pro Leu Thr Ser Ser Asn His Ser Thr Ser Pro Gln Leu Ser Thr				
1025		1030		1035
Gly Val Ser Phe Phe Phe Leu Ser Phe His Ile Ser Asn Leu Gln				
1040		1045		1050
Phe Asn Ser Ser Leu Glu Asp Pro Ser Thr Asp Tyr Tyr Gln Glu				
1055		1060		1065
Leu Gln Arg Asp Ile Ser Glu Met Phe Leu Gln Ile Tyr Lys Gln				
1070		1075		1080
Gly Gly Phe Leu Gly Leu Ser Asn Ile Lys Phe Arg Pro Gly Ser				
1085		1090		1095
Val Val Val Gln Leu Thr Leu Ala Phe Arg Glu Gly Thr Ile Asn				
1100		1105		1110
Val His Asp Val Glu Thr Gln Phe Asn Gln Tyr Lys Thr Glu Ala				
1115		1120		1125
Ala Ser Arg Tyr Asn Leu Thr Ile Ser Asp Val Ser Val Ser Asp				
1130		1135		1140
Val Pro Phe Pro Phe Ser Ala Gln Ser Gly Ala Gly Val Pro Gly				
1145		1150		1155
Trp Gly Ile Ala Leu Leu Val Leu Val Cys Val Leu Val Ala Leu				
1160		1165		1170
Ala Ile Val Tyr Leu Ile Ala Leu Ala Val Cys Gln Cys Arg Arg				
1175		1180		1185
Lys Asn Tyr Gly Gln Leu Asp Ile Phe Pro Ala Arg Asp Thr Tyr				
1190		1195		1200
His Pro Met Ser Glu Tyr Pro Thr Tyr His Thr His Gly Arg Tyr				
1205		1210		1215
Val Pro Pro Ser Ser Thr Asp Arg Ser Pro Tyr Glu Lys Val Ser				
1220		1225		1230
Ala Gly Asn Gly Gly Ser Ser Leu Ser Tyr Thr Asn Pro Ala Val				
1235		1240		1245
Ala Ala Ala Ser Ala Asn Leu				
1250		1255		

<212> PRT  
<213> Artificial Sequence

<220>  
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<400> 2

Gly Thr Ile Asn Val His Asp Val Glu Thr Gln Phe Asn Gln Tyr Lys  
1 5 10 15

Thr Glu Ala Ala Ser Arg Tyr Asn Leu Thr Ile Ser Asp Val Ser Val  
20 25 30

Ser Asp Val Pro Phe Pro Phe Ser Ala Gln Ser Gly Ala  
35 40 45

<210> 3  
<211> 35  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> N-10 peptide

<400> 3

Gln Phe Asn Gln Tyr Lys Thr Glu Ala Ala Ser Arg Tyr Asn Leu Thr  
1 5 10 15

Ile Ser Asp Val Ser Val Ser Asp Val Pro Phe Pro Phe Ser Ala Gln  
20 25 30

Ser Gly Ala  
35

<210> 4  
<211> 26  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> N-19 peptide

<400> 4

Ala Ser Arg Tyr Asn Leu Thr Ile Ser Asp Val Ser Val Ser Asp Val  
1 5 10 15

Pro Phe Pro Phe Ser Ala Gln Ser Gly Ala  
20 25

<210> 5  
<211> 22  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> N-23 peptide

<400> 5

Asn Leu Thr Ile Ser Asp Val Ser Val Ser Asp Val Pro Phe Pro Phe  
1 5 10 15

Ser Ala Gln Ser Gly Ala  
20

<210> 6  
<211> 19  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> N-26 peptide

<400> 6

Ile Ser Asp Val Ser Val Ser Asp Val Pro Phe Pro Phe Ser Ala Gln  
1 5 10 15

Ser Gly Ala

<210> 7  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> N-30 peptide

<400> 7

Ser Val Ser Asp Val Pro Phe Pro Phe Ser Ala Gln Ser Gly Ala  
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<210> 8  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> N-10/C-5 peptide

<400> 8

Gln Phe Asn Gln Tyr Lys Thr Glu Ala Ala Ser Arg Tyr Asn Leu Thr  
1 5 10 15

Ile Ser Asp Val Ser Val Ser Asp Val Pro Phe Pro Phe Ser  
20 25 30

<210> 9  
<211> 21  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> N-19/C-5 peptide

<400> 9

Ala Ser Arg Tyr Asn Leu Thr Ile Ser Asp Val Ser Val Ser Asp Val  
1 5 10 15

Pro Phe Pro Phe Ser  
20

<210> 10  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> N-36 peptide

<400> 10

Phe Pro Phe Ser Ala Gln Ser Gly Ala  
1 5

<210> 11

<400> 11  
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<210> 12  
<211> 366  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 Heavy chain variable region sequence

<400> 12  
gaggatgaagg tggatggagtc tgggggagac ttagtgaagc ctggagggtc cctgaaactc 60  
tcctgtgtag tctctggatt cactttcagt agatatggca tgtcttgggt tcgccagact 120  
ccaggcaaga ggctggagtg ggtcgcaacc attagtgggt gcggtactta catctactat 180  
ccagacagtg tgaaggggcg attcaccatc tccagagaca atgccaagaa caccctgtac 240  
ctgcaaata gaagtctgaa gtctgaggac acagccatgt atcactgtac aagggataac 300  
tacggttagga actacgacta cggatatggac tactgggggtc aaggaacctc agtcaccgtc 360  
tcctca 366

<210> 13  
<211> 122  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 Heavy chain variable region sequence

<400> 13

Glu Val Lys Val Val Glu Ser Gly Gly Asp Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Val Val Ser Gly Phe Thr Phe Ser Arg Tyr  
20 25 30

Gly Met Ser Trp Val Arg Gln Thr Pro Gly Lys Arg Leu Glu Trp Val  
35 40 45

Ala Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr  
65 70 75 80

Leu Gln Met Ser Ser Leu Lys Ser Glu Asp Thr Ala Met Tyr His Cys  
85 90 95

Thr Arg Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr Gly Met Asp Tyr Trp  
100 105 110

Gly Gln Gly Thr Ser Val Thr Val Ser Ser  
115 120

<210> 14  
<211> 75  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable framework region 1 (FWR1) sequence

<400> 14  
gaggtgaagg tgggtggagtc tgggggagac ttagtgaagc ctggagggtc cctgaaactc 60  
tcctgtgtag tctct 75

<210> 15  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable framework region 1 (FWR1) sequence

<400> 15

Glu Val Lys Val Val Glu Ser Gly Gly Asp Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Val Val Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 16  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 16  
ggattcactt tcagtagata tggcatgtct 30

<210> 17  
<211> 5  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse E6 heavy chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 17

Arg Tyr Gly Met Ser  
1 5

<210> 18  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable framework region 2 (FWR2) sequence

<400> 18  
tgggttcgcc agactccagg caagaggctg gagtgggtcg ca 42

<210> 19  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable framework region 2 (FWR2) sequence

<400> 19

Trp Val Arg Gln Thr Pro Gly Lys Arg Leu Glu Trp Val Ala  
1 5 10

<210> 20  
<211> 51  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 20  
accattagtg gtggcggtac ttacatctac tatccagaca gtgtgaaggg g 51

<210> 21  
<211> 17  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 21

Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys  
1 5 10 15

Gly

<210> 22  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable framework region 3 (FWR3) acid  
sequence

<400> 22  
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aagtctgagg acacagccat gtatcactgt acaagg 96

<210> 23  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable framework region 3 (FWR3) acid  
sequence

<400> 23

Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln  
1 5 10 15

Met Ser Ser Leu Lys Ser Glu Asp Thr Ala Met Tyr His Cys Thr Arg  
20 25 30

<210> 24  
<211> 39  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable complementarity determining regions  
3 (CDR3) sequence:

<400> 24  
gataactacg gtaggaacta cgactacggt atggactac 39

<210> 25  
<211> 13  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 heavy chain variable complementarity determining regions  
3 (CDR3) sequence:

<400> 25

Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr Gly Met Asp Tyr  
1 5 10

<210> 26

<400> 26



000

<210> 27

<400> 27  
000

<210> 28

<400> 28  
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<400> 29  
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<210> 37

<400> 37  
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<210> 38

<211> 366

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized E6 heavy chain variable region sequence from  
IGHV3-21\*03

<400> 38

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<210> 41  
<211> 30  
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<223> Humanized E6 heavy chain variable framework region 1 (FWR1) acid  
sequence

<400> 41

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 42  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized E6 heavy chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 42  
aggtatggca tgagc 15

<210> 43  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized E6 heavy chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 43

Arg Tyr Gly Met Ser  
1 5

<210> 44  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized E6 heavy chain variable framework region 2 (FWR2) acid  
sequence

<400> 44  
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<210> 45  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized E6 heavy chain variable framework region 2 (FWR2) acid

sequence

<400> 45

Trp Val Arg Gln Ala Pro Gly Lys Arg Leu Glu Trp Val Ser  
1 5 10

<210> 46

<211> 51

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized E6 heavy chain variable complementarity determining  
regions 2 (CDR2) sequence

<400> 46

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<210> 47

<211> 17

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized E6 heavy chain variable complementarity determining  
regions 2 (CDR2) sequence

<400> 47

Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys  
1 5 10 15

Gly

<210> 48

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized E6 heavy chain variable framework region 3 (FWR3) acid  
sequence

<400> 48

cgattcacca tctccagaga caacgccaaag aacaccctgt atctgcaaat gaacagcctg 60

agagccgagg acacggctgt gtattactgt accaga 96

<210> 49

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized E6 heavy chain variable framework region 3 (FWR3) acid  
sequence

<400> 49

Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln  
1 5 10 15

Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys Thr Arg  
20 25 30

<210> 50  
<211> 39  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized E6 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 50  
gataactatg gccgcaacta tgattatggc atggattat 39

<210> 51  
<211> 13  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized E6 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 51

Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr Gly Met Asp Tyr  
1 5 10

<210> 52  
<211> 1442  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized E6 IgG2 heavy chain synthesized by Genescript

<400> 52  
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ttagaagggtg tccagtgtga ggtgcagctg gtggagtctg ggggaggcct ggtcaagcct 120  
gggggggtccc tgagactctc ctgtgcagcc tctggattca ccttcagtag gtatggcatg 180  
agctgggtcc gccaggctcc aggggaagagg ctggagtggg tctcaaccat tagtggcgga 240  
ggcacctaca tatactaccc agactcagtg aagggccgat tcaccatctc cagagacaac 300  
gccaagaaca ccctgtatct gcaaatgaac agcctgagag ccgaggacac ggctgtgtat 360  
tactgtacca gagataacta tggccgcaac tatgattatg gcatggatta ttggggccag 420  
ggcaccctgg tgaccgtgag cagcgcctcc accaagggcc catcggtctt cccctggcg 480  
ccctgctcca ggagcacctc cgagagcaca gccgccctgg gctgcctggg caaggactac 540  
ttccccgaac cggtgacggt gtcgtggaac tcaggcgctc tgaccagcgg cgtgcacacc 600  
ttcccagctg tcctacagtc ctcaggactc tactccctca gcagcgtggg gaccgtgccc 660  
tccagcaact tcggcaccca gacctacacc tgcaacgtag atcacaagcc cagcaacacc 720  
aaggtggaca agacagttga gcgcaaatgt tgtgtcgagt gccaccgtg cccagcacca 780  
cctgtggcag gaccgtcagt cttcctcttc ccccaaaaac ccaaggacac cctcatgatc 840

tcccggaccc ctgaggtcac gtgcgtggtg gtggacgtga gccacgaaga ccccgaggtc	900
cagttcaact ggtacgtgga cggcgtggag gtgcataatg ccaagacaaa gccacgggag	960
gagcagttca acagcacgtt ccgtgtggtc agcgtcctca ccgttgtgca ccaggactgg	1020
ctgaacggca aggagtacaa gtgcaaggtc tccaacaaag gcctcccagc ccccatcgag	1080
aaaaccatct ccaaaaccaa agggcagccc cgagaaccac aggtgtacac cctgccccca	1140
tcccgggagg agatgaccaa gaaccaggtc agcctgacct gcctgggtcaa aggcttctac	1200
cccagcgaca tcgccgtgga gtgggagagc aatgggcagc cggagaacaa ctacaagacc	1260
acacctccca tgctggactc cgacggctcc ttcttcctct acagcaagct caccgtggac	1320
aagagcaggt ggacgaggg gaacgtcttc tcatgctccg tgatgcatga ggctctgcac	1380
aaccactaca cgcagaagag cctctccctg tctccgggta aatagtaagt ttaaactcta	1440
ga	1442

<210> 53  
 <211> 477  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Humanized E6 IgG2 heavy chain synthesized by Genescript

<220>  
 <221> misc\_feature  
 <222> (477)..(477)  
 <223> Xaa can be any naturally occurring amino acid

<400> 53

Glu	Phe	Ala	Trp	Ala	Thr	Met	Glu	Leu	Gly	Leu	Arg	Trp	Val	Phe	Leu
1				5					10					15	

Val	Ala	Ile	Leu	Glu	Gly	Val	Gln	Cys	Glu	Val	Gln	Leu	Val	Glu	Ser
			20					25					30		

Gly	Gly	Gly	Leu	Val	Lys	Pro	Gly	Gly	Ser	Leu	Arg	Leu	Ser	Cys	Ala
		35					40					45			

Ala	Ser	Gly	Phe	Thr	Phe	Ser	Arg	Tyr	Gly	Met	Ser	Trp	Val	Arg	Gln
	50					55				60					

Ala	Pro	Gly	Lys	Arg	Leu	Glu	Trp	Val	Ser	Thr	Ile	Ser	Gly	Gly	Gly
65					70				75					80	

Thr	Tyr	Ile	Tyr	Tyr	Pro	Asp	Ser	Val	Lys	Gly	Arg	Phe	Thr	Ile	Ser
			85						90					95	

Arg	Asp	Asn	Ala	Lys	Asn	Thr	Leu	Tyr	Leu	Gln	Met	Asn	Ser	Leu	Arg
			100					105						110	

Ala	Glu	Asp	Thr	Ala	Val	Tyr	Tyr	Cys	Thr	Arg	Asp	Asn	Tyr	Gly	Arg
-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----



Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn  
405 410 415

Tyr Lys Thr Thr Pro Pro Met Leu Asp Ser Asp Gly Ser Phe Phe Leu  
420 425 430

Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val  
435 440 445

Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr Thr Gln  
450 455 460

Lys Ser Leu Ser Leu Ser Pro Gly Lys Val Thr Leu Xaa  
465 470 475

<210> 54  
<211> 981  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Human IgG2 heavy chain constant region sequence

<400> 54  
gcctccacca agggcccatc ggtcttcccc ctggcgcctt gctccaggag cacctccgag 60  
agcacagccg ccctgggctg cctggtcaag gactacttcc ccgaaccggg gacgggtgtcg 120  
tggaactcag gcgctctgac cagcggcgtg cacaccttcc cagctgtcct acagtcctca 180  
ggactctact ccctcagcag cgtggtgacc gtgccctcca gcaacttcgg caccagacc 240  
tacacctgca acgtagatca caagcccagc aacaccaagg tggacaagac agttgagcgc 300  
aaatgttgtg tcgagtgtcc accgtgtcca gcaccacctg tggcaggacc gtcagtcttc 360  
ctcttcccc caaaaccaa ggacaccctc atgatctccc ggaccctga ggtcacgtgc 420  
gtggtggtgg acgtgagcca cgaagacccc gaggtccagt tcaactggta cgtggacggc 480  
gtggagggtgc ataatgcaa gacaaagcca cgggaggagc agttcaacag cacgttccgt 540  
gtggtcagcg tcctcaccgt tgtgcaccag gactggctga acggcaagga gtacaagtgc 600  
aaggtctcca acaaaggcct ccagccccc atcgagaaaa ccatctcaa aaccaaaggg 660  
cagccccgag aaccacaggt gtacaccctg ccccatccc gggaggagat gaccaagaac 720  
caggtcagcc tgacctgcct ggtcaaaggc ttctaccca gcgacatcgc cgtggagtgg 780  
gagagcaatg ggagccgga gaacaactac aagaccacac ctccatgct ggactccgac 840  
ggctccttct tcctctacag caagctcacc gtggacaaga gcaggtggca gcagggaac 900  
gtcttctcat gctccgtgat gcatgaggct ctgcacaacc actacacgca gaagagcctc 960  
tccctgtctc cgggtaaata g 981

<210> 55  
<211> 326  
<212> PRT



<213> Artificial Sequence

<220>

<223> Human IgG2 heavy chain constant region sequence

<400> 55

Ala Ser Thr Lys Gly Pro Ser Val Phe Pro Leu Ala Pro Cys Ser Arg  
1 5 10 15

Ser Thr Ser Glu Ser Thr Ala Ala Leu Gly Cys Leu Val Lys Asp Tyr  
20 25 30

Phe Pro Glu Pro Val Thr Val Ser Trp Asn Ser Gly Ala Leu Thr Ser  
35 40 45

Gly Val His Thr Phe Pro Ala Val Leu Gln Ser Ser Gly Leu Tyr Ser  
50 55 60

Leu Ser Ser Val Val Thr Val Pro Ser Ser Asn Phe Gly Thr Gln Thr  
65 70 75 80

Tyr Thr Cys Asn Val Asp His Lys Pro Ser Asn Thr Lys Val Asp Lys  
85 90 95

Thr Val Glu Arg Lys Cys Cys Val Glu Cys Pro Pro Cys Pro Ala Pro  
100 105 110

Pro Val Ala Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp  
115 120 125

Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp  
130 135 140

Val Ser His Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr Val Asp Gly  
145 150 155 160

Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Phe Asn  
165 170 175

Ser Thr Phe Arg Val Val Ser Val Leu Thr Val Val His Gln Asp Trp  
180 185 190

Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly Leu Pro  
195 200 205

Ala Pro Ile Glu Lys Thr Ile Ser Lys Thr Lys Gly Gln Pro Arg Glu  
210 215 220

Pro Gln Val Tyr Thr Leu Pro Pro Ser Arg Glu Glu Met Thr Lys Asn  
225 230 235 240

Gln Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile  
245 250 255

Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr  
260 265 270

Thr Pro Pro Met Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys  
275 280 285

Leu Thr Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys  
290 295 300

Ser Val Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu  
305 310 315 320

Ser Leu Ser Pro Gly Lys  
325

<210> 56  
<211> 1362  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized E6 IgG1 heavy chain sequence

<400> 56  
gaggtgcagc tggaggagtc tgggggaggc ctggtcaagc ctggggggtc cctgagactc 60  
tcctgtgcag cctctggatt caccttcagt aggtatggca tgagctgggt ccgccaggct 120  
ccagggaaga ggctggagtg ggtctcaacc attagtggcg gaggcaccta catatactac 180  
ccagactcag tgaagggccg attcaccatc tccagagaca acgccaagaa cccactgtat 240  
ctgcaaatga acagcctgag agccgaggac acggctgtgt attactgtcc cagagataac 300  
tatggccgca actatgatta tggcatggat tattggggcc agggcaccct ggtgaccgtg 360  
agcagcgcta gcaccaaggg cccatcggtc tttcccctgg caccctcctc caagagcacc 420  
tctgggggca cagcggccct gggctgcctg gtcaaggact acttccccga accggtgacg 480  
gtgtcgtgga actcaggcgc cctgaccagc ggcgtgcaca ctttcccggc tgtcctacag 540  
tcctcaggac tctactccct cagcagcgtg gtgacagtgc cctccagcag cttgggcacc 600  
cagacctaca tctgcaacgt gaatcacaag cccagcaaca ccaaggtgga caagaaagt 660  
gagcccaaat cttgtgacaa aactcacaca tgcccaccgt gcccagcacc tgaactcctg 720  
gggggaccgt cagtcttcct cttccccca aaaccaagg acaccctcat gatctcccgg 780  
accctgagg tcacatgcgt ggtggtggac gtgagccacg aagaccctga ggtcaagttc 840  
aactggtacg tggacggcgt ggaggtgcat aatgccaaga caaagccgcg ggaggagcag 900  
tacaacagca cgtaccgtgt ggtcagcgtc ctcaccgtcc tgcaccagga ctggctgaat 960  
ggcaaggagt acaagtgcaa ggtctccaac aaagccctcc cagccccat cgagaaaacc 1020  
atctccaaag ccaaagggca gccccgagaa ccacaggtgt acaccctgcc cccatcccgg 1080  
gaggagatga ccaagaacca ggtcagcctg acctgcctgg tcaaaggctt ctatcccagc 1140

gacatcgccg tggagtggga gagcaatggg cagccggaga acaactacaa gaccacgcct	1200
cccggtgctgg actccgacgg ctcttcttctc ctctacagca agctcaccgt ggacaagagc	1260
aggtggcagc aggggaacgt cttctcatgc tccgtgatgc atgaggctct gcacaaccac	1320
tacacgcaga agagcctctc cctgtctccg ggtaaataat aa	1362

<210> 57  
 <211> 452  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Humanized E6 IgG1 heavy chain sequence

<400> 57

Glu	Val	Gln	Leu	Val	Glu	Ser	Gly	Gly	Gly	Leu	Val	Lys	Pro	Gly	Gly
1				5					10					15	

Ser	Leu	Arg	Leu	Ser	Cys	Ala	Ala	Ser	Gly	Phe	Thr	Phe	Ser	Arg	Tyr
			20					25					30		

Gly	Met	Ser	Trp	Val	Arg	Gln	Ala	Pro	Gly	Lys	Arg	Leu	Glu	Trp	Val
	35						40					45			

Ser	Thr	Ile	Ser	Gly	Gly	Gly	Thr	Tyr	Ile	Tyr	Tyr	Pro	Asp	Ser	Val
	50					55					60				

Lys	Gly	Arg	Phe	Thr	Ile	Ser	Arg	Asp	Asn	Ala	Lys	Asn	Pro	Leu	Tyr
65					70					75				80	

Leu	Gln	Met	Asn	Ser	Leu	Arg	Ala	Glu	Asp	Thr	Ala	Val	Tyr	Tyr	Cys
			85						90					95	

Pro	Arg	Asp	Asn	Tyr	Gly	Arg	Asn	Tyr	Asp	Tyr	Gly	Met	Asp	Tyr	Trp
			100					105					110		

Gly	Gln	Gly	Thr	Leu	Val	Thr	Val	Ser	Ser	Ala	Ser	Thr	Lys	Gly	Pro
		115					120					125			

Ser	Val	Phe	Pro	Leu	Ala	Pro	Ser	Ser	Lys	Ser	Thr	Ser	Gly	Gly	Thr
	130					135					140				

Ala	Ala	Leu	Gly	Cys	Leu	Val	Lys	Asp	Tyr	Phe	Pro	Glu	Pro	Val	Thr
145					150					155				160	

Val	Ser	Trp	Asn	Ser	Gly	Ala	Leu	Thr	Ser	Gly	Val	His	Thr	Phe	Pro
			165						170					175	

Ala	Val	Leu	Gln	Ser	Ser	Gly	Leu	Tyr	Ser	Leu	Ser	Ser	Val	Val	Thr
		180					185						190		

Val	Pro	Ser	Ser	Ser	Leu	Gly	Thr	Gln	Thr	Tyr	Ile	Cys	Asn	Val	Asn
		195					200					205			

His Lys Pro Ser Asn Thr Lys Val Asp Lys Lys Val Glu Pro Lys Ser  
 210 215 220  
 Cys Asp Lys Thr His Thr Cys Pro Pro Cys Pro Ala Pro Glu Leu Leu  
 225 230 235 240  
 Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu  
 245 250 255  
 Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser  
 260 265 270  
 His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val Asp Gly Val Glu  
 275 280 285  
 Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn Ser Thr  
 290 295 300  
 Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu Asn  
 305 310 315 320  
 Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Ala Leu Pro Ala Pro  
 325 330 335  
 Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln  
 340 345 350  
 Val Tyr Thr Leu Pro Pro Ser Arg Glu Glu Met Thr Lys Asn Gln Val  
 355 360 365  
 Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val  
 370 375 380  
 Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro  
 385 390 395 400  
 Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr  
 405 410 415  
 Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val  
 420 425 430  
 Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu  
 435 440 445  
 Ser Pro Gly Lys  
 450

<210> 58  
 <211> 996  
 <212> DNA

<213> Artificial Sequence

<220>

<223> Human IgG1 heavy chain constant region sequence

<400> 58

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gctagcacca agggcccatc ggtcttcccc ctggcaccct cctccaagag cacctctggg      60
ggcacagcgg ccctgggctg cctgggtcaag gactacttcc ccgaaccggt gacggtgtcg      120
tggaactcag gcgccctgac cagcggcgtg cacaccttcc cggctgtcct acagtcctca      180
ggactctact ccctcagcag cgtgggtgaca gtgccctcca gcagcttggg caccagacc      240
tacatctgca acgtgaatca caagcccagc aacaccaagg tggacaagaa agttgagccc      300
aaatcttgtg acaaaactca cacatgcccc ccgtgcccag cacctgaact cctgggggga      360
ccgtcagtct tcctcttccc cccaaaaccc aaggacaccc tcatgatctc ccggaccct      420
gaggtcacat gcgtgggtgt ggacgtgagc cacgaagacc ctgaggtcaa gttcaactgg      480
tacgtggacg gcgtggaggt gcataatgcc aagacaaagc cgcgggagga gcagtacaac      540
agcacgtacc gtgtgggtcag cgtcctcacc gtcctgcacc aggactggct gaatggcaag      600
gagtacaagt gcaaggtctc caacaaagcc ctcccagccc ccatcgagaa aaccatctcc      660
aaagccaaag ggagccccg agaaccacag gtgtacaccc tgccccatc ccgggaggag      720
atgaccaaga accaggtcag cctgacctgc ctgggtcaaag gcttctatcc cagcgacatc      780
gccgtggagt gggagagcaa tgggcagccg gagaacaact acaagaccac gcctcccgtg      840
ctggactccg acggctcctt ctctctctac agcaagctca ccgtggacaa gagcaggtgg      900
cagcagggga acgtcttctc atgctccgtg atgcatgagg ctctgcacaa ccactacacg      960
cagaagagcc tctccctgtc tccgggtaaa tgataa      996
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<210> 59

<211> 330

<212> PRT

<213> Artificial Sequence

<220>

<223> Human IgG1 heavy chain constant region sequence

<400> 59

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Ala Ser Thr Lys Gly Pro Ser Val Phe Pro Leu Ala Pro Ser Ser Lys
1           5           10          15

Ser Thr Ser Gly Gly Thr Ala Ala Leu Gly Cys Leu Val Lys Asp Tyr
          20          25          30

Phe Pro Glu Pro Val Thr Val Ser Trp Asn Ser Gly Ala Leu Thr Ser
          35          40          45

Gly Val His Thr Phe Pro Ala Val Leu Gln Ser Ser Gly Leu Tyr Ser
50          55          60

Leu Ser Ser Val Val Thr Val Pro Ser Ser Ser Leu Gly Thr Gln Thr
65          70          75          80
```

Tyr Ile Cys Asn Val Asn His Lys Pro Ser Asn Thr Lys Val Asp Lys  
 85 90 95

Lys Val Glu Pro Lys Ser Cys Asp Lys Thr His Thr Cys Pro Pro Cys  
 100 105 110

Pro Ala Pro Glu Leu Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro  
 115 120 125

Lys Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys  
 130 135 140

Val Val Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp  
 145 150 155 160

Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu  
 165 170 175

Glu Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu  
 180 185 190

His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn  
 195 200 205

Lys Ala Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly  
 210 215 220

Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Arg Glu Glu  
 225 230 235 240

Met Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr  
 245 250 255

Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn  
 260 265 270

Asn Tyr Lys Thr Thr Pro Pro Val Leu Asp Ser Asp Gly Ser Phe Phe  
 275 280 285

Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln Gln Gly Asn  
 290 295 300

Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr Thr  
 305 310 315 320

Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys  
 325 330

<210> 60

<400> 60

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<210> 61

<400> 61  
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<210> 62

<400> 62  
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<210> 63

<400> 63  
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<210> 64

<400> 64  
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<210> 65

<211> 318

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse E6 Light Chain variable region sequence

<400> 65

caaattgttc tcacccagtc tccagcaatc atgtctgcat ctccagggga ggaggtcacc 60

ctaacctgca gtgccacctc aagtgttaagt tacatacact ggttccagca gaggccaggc 120

acttctccca aactctggat ttatagcaca tccaacctgg cttctggagt ccctgttcgc 180

ttcagtggca gtggatatgg gacctcttac tctctcacia tcagccgaat ggaggctgaa 240

gatgctgcca cttattactg ccagcaaagg agtagttccc cattcacgtt cggctcgggg 300

acaaagttgg aaataaaa 318

<210> 66

<211> 106

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse E6 Light Chain variable region sequence

<400> 66

Gln Ile Val Leu Thr Gln Ser Pro Ala Ile Met Ser Ala Ser Pro Gly  
1 5 10 15

Glu Glu Val Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile  
20 25 30

His Trp Phe Gln Gln Arg Pro Gly Thr Ser Pro Lys Leu Trp Ile Tyr  
35 40 45

Ser Thr Ser Asn Leu Ala Ser Gly Val Pro Val Arg Phe Ser Gly Ser  
50 55 60

Gly Tyr Gly Thr Ser Tyr Ser Leu Thr Ile Ser Arg Met Glu Ala Glu  
65 70 75 80

Asp Ala Ala Thr Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr  
85 90 95

Phe Gly Ser Gly Thr Lys Leu Glu Ile Lys  
100 105

<210> 67  
<211> 69  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable framework region 1 (FWR1) sequence

<400> 67  
caaattgttc tcaccagtc tccagcaatc atgtctgcat ctccagggga ggaggtcacc 60  
ctaacctgc 69

<210> 68  
<211> 23  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable framework region 1 (FWR1) sequence

<400> 68

Gln Ile Val Leu Thr Gln Ser Pro Ala Ile Met Ser Ala Ser Pro Gly  
1 5 10 15

Glu Glu Val Thr Leu Thr Cys  
20

<210> 69  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 69  
agtgccacct caagtgtgtaag ttacatacac 30

<210> 70  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 70



Ser Ala Thr Ser Ser Val Ser Tyr Ile His  
1 5 10

<210> 71  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable framework region 2 (FWR2) sequence

<400> 71  
tggttcagc agaggccagg cacttctccc aaactctgga tttat 45

<210> 72  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable framework region 2 (FWR2) sequence

<400> 72

Trp Phe Gln Gln Arg Pro Gly Thr Ser Pro Lys Leu Trp Ile Tyr  
1 5 10 15

<210> 73  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 73  
agcacatcca acctggcttc t 21

<210> 74  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 74

Ser Thr Ser Asn Leu Ala Ser  
1 5

<210> 75  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable framework region 3 (FWR3) sequence

<400> 75  
ggagtcctg ttcgcttcag tggcagtgga tatgggacct cttactctct cacaatcagc 60

cgaatggagg ctgaagatgc tgccacttat tactgc

96

<210> 76  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable framework region 3 (FWR3) sequence

<400> 76

Gly	Val	Pro	Val	Arg	Phe	Ser	Gly	Ser	Gly	Tyr	Gly	Thr	Ser	Tyr	Ser
1				5					10					15	

Leu	Thr	Ile	Ser	Arg	Met	Glu	Ala	Glu	Asp	Ala	Ala	Thr	Tyr	Tyr	Cys
			20					25					30		

<210> 77  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 77  
cagcaaagga gtagttcccc attcacg

27

<210> 78  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse E6 light chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 78

Gln	Gln	Arg	Ser	Ser	Ser	Pro	Phe	Thr
1			5					

<210> 79

<400> 79  
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<210> 80

<400> 80  
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<210> 81

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<210> 91

<400> 91  
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<210> 92

<400> 92  
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<210> 93

<211> 318

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized E6 light chain variable region sequence from  
IGKV3-11\*02

<400> 93

gaaattgtgt tgacacagtc tccagccacc ctgtctttgt ctccagggga aagagccacc 60

ctcacctgca gcgccaccag cagtgttagc tacatccact ggtaccaaca gaggcctggc 120

cagagcccca ggctcctcat ctatagcacc tccaacctgg ccagcggcat cccagccagg 180

ttcagtggca gtgggtctgg gagcgactac actctcacca tcagcagcct agagcctgaa 240

gattttgcag tttattactg tcagcagcgt agcagctccc ctttcacctt tggcagcggc 300

<210> 94  
<211> 106  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized E6 light chain variable region sequence from  
IGKV3-11\*02

<400> 94

Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly  
1 5 10 15

Glu Arg Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile  
20 25 30

His Trp Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr  
35 40 45

Ser Thr Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser  
50 55 60

Gly Ser Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu  
65 70 75 80

Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr  
85 90 95

Phe Gly Ser Gly Thr Lys Val Glu Ile Lys  
100 105

<210> 95  
<211> 69  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized E6 light chain variable framework region 1 (FWR1) acid  
sequence

<400> 95  
gaaattgtgt tgacacagtc tccagccacc ctgtctttgt ctccagggga aagagccacc 60  
ctcacctgc 69

<210> 96  
<211> 23  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized E6 light chain variable framework region 1 (FWR1) acid  
sequence

<400> 96

Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly  
1 5 10 15

Glu Arg Ala Thr Leu Thr Cys  
20

<210> 97  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized E6 light chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 97  
agcgccacca gcagtgttag ctacatccac 30

<210> 98  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized E6 light chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 98

Ser Ala Thr Ser Ser Val Ser Tyr Ile His  
1 5 10

<210> 99  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized E6 heavy light variable framework region 2 (FWR2) acid  
sequence

<400> 99  
tggtaccaac agaggcctgg ccagagcccc aggtctctca tctat 45

<210> 100  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized E6 heavy light variable framework region 2 (FWR2) acid  
sequence

<400> 100

Trp Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr  
1 5 10 15

<210> 101  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
 <223> Humanized E6 light chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 101  
 agcacctcca acctggccag c 21  
  
 <210> 102  
 <211> 7  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized E6 light chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 102  
  
 Ser Thr Ser Asn Leu Ala Ser  
 1 5  
  
 <210> 103  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized E6 light chain variable framework region 3 (FWR3) acid sequence  
  
 <400> 103  
 ggcatcccag ccaggttcag tggcagtggg tctgggagcg actacactct caccatcagc 60  
 agcctagagc ctgaagattt tgcagtttat tactgt 96  
  
 <210> 104  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized E6 light chain variable framework region 3 (FWR3) acid sequence  
  
 <400> 104  
  
 Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Ser Asp Tyr Thr  
 1 5 10 15  
  
 Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys  
 20 25 30  
  
 <210> 105  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized E6 light chain variable complementarity determining regions 3 (CDR3) sequence  
  
 <400> 105  
 cagcagcgta gcagctcccc ttccacc 27

<210> 106  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized E6 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 106

Gln Gln Arg Ser Ser Ser Pro Phe Thr  
 1 5

<210> 107  
 <211> 740  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized E6 Kappa light chain synthesized by Genescript

<400> 107  
 gaattctaag cttgggccac catggaagcc ccagcgcagc ttctcttcct cctgctactc 60  
 tggctcccag ataccactgg agaaattgtg ttgacacagt ctccagccac cctgtctttg 120  
 tctccagggg aaagagccac cctcacctgc agcggcacca gcagtgttag ctacatccac 180  
 tggtagcaac agaggcctgg ccagagcccc aggctcctca tctatagcac ctccaacctg 240  
 gccagcggca tcccagccag gttcagtggc agtgggtctg ggagcgacta cactctcacc 300  
 atcagcagcc tagagcctga agattttgca gtttattact gtcagcagcg tagcagctcc 360  
 cctttcacct ttggcagcgg caccaaagtg gaaattaaaa ggacggtggc tgcaccatct 420  
 gtcttcatct tccgccatc tgatgagcag ttgaaatctg gaactgcctc tgttgtgtgc 480  
 ctgctgaata acttctatcc cagagaggcc aaagtacagt ggaagggtgga taacgccctc 540  
 caatcgggta actcccagga gagtgtcaca gagcaggaca gcaaggacag cacctacagc 600  
 ctcagcagca ccctgacgct gagcaaagca gactacgaga aacacaaaagt ctacgcctgc 660  
 gaagtcaccc atcagggcct gagctcgccc gtcacaaaga gcttcaacag gggagagtgt 720  
 tagtaagttt aaactctaga 740

<210> 108  
 <211> 243  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized E6 Kappa light chain synthesized by Genescript

<220>  
 <221> misc\_feature  
 <222> (243)..(243)  
 <223> Xaa can be any naturally occurring amino acid

<400> 108

Glu Phe Ala Trp Ala Thr Met Glu Ala Pro Ala Gln Leu Leu Phe Leu

1	5	10	15
Leu Leu Leu Trp 20	Leu Pro Asp Thr Thr 25	Gly Glu Ile Val 30	Leu Thr Gln
Ser Pro Ala Thr 35	Leu Ser Leu Ser 40	Pro Gly Glu Arg 45	Ala Thr Leu Thr
Cys Ser Ala Thr 50	Ser Ser Val Ser 55	Tyr Ile His Trp 60	Tyr Gln Gln Arg
Pro Gly Gln Ser 65	Pro Arg Leu Leu 70	Ile Tyr Ser Thr 75	Ser Asn Leu Ala 80
Ser Gly Ile Pro 85	Ala Arg Phe Ser 90	Gly Ser Gly Ser 95	Gly Ser Asp Tyr
Thr Leu Thr Ile 100	Ser Ser Leu Glu 105	Pro Glu Asp Phe 110	Ala Val Tyr Tyr
Cys Gln Gln Arg 115	Ser Ser Ser Pro 120	Phe Thr Phe Gly 125	Ser Gly Thr Lys
Val Glu Ile Lys 130	Arg Thr Val Ala 135	Ala Pro Ser Val 140	Phe Ile Phe Pro
Pro Ser Asp Glu 145	Gln Leu Lys Ser 150	Gly Thr Ala Ser 155	Val Val Cys Leu 160
Leu Asn Asn Phe 165	Tyr Pro Arg Glu 170	Ala Lys Val Gln 175	Trp Lys Val Asp
Asn Ala Leu Gln 180	Ser Gly Asn Ser 185	Gln Glu Ser Val 190	Thr Glu Gln Asp
Ser Lys Asp Ser 195	Thr Tyr Ser Leu 200	Ser Ser Thr Leu 205	Thr Leu Ser Lys
Ala Asp Tyr Glu 210	Lys His Lys Val 215	Tyr Ala Cys Glu 220	Val Thr His Gln
Gly Leu Ser Ser 225	Pro Val Thr Lys 230	Ser Phe Asn Arg 235	Gly Glu Cys Val 240
Thr Leu Xaa			

<210> 109  
 <211> 324  
 <212> DNA  
 <213> Artificial Sequence  
 <220>



<223> Human Kappa light chain constant region sequence

<400> 109

```
aggacggtgg ctgcaccatc tgtcttcac ttcccgccat ctgatgagca gttgaaatct      60
ggaactgcct ctgttggttg cctgctgaat aacttctatc ccagagaggc caaagtacag      120
tggaaggtgg ataacgccct ccaatcgggt aactcccagg agagtgtcac agagcaggac      180
agcaaggaca gcacctacag cctcagcagc accctgacgc tgagcaaagc agactacgag      240
aaacacaaag tctacgcctg cgaagtcacc catcagggcc tgagctcgcc cgtcacaaag      300
agcttcaaca ggggagagtg ttag                                             324
```

<210> 110

<211> 107

<212> PRT

<213> Artificial Sequence

<220>

<223> Human Kappa light chain constant region sequence

<400> 110

```
Arg Thr Val Ala Ala Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu
1              5              10              15
```

```
Gln Leu Lys Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe
                20              25              30
```

```
Tyr Pro Arg Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln
                35              40              45
```

```
Ser Gly Asn Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser
50              55              60
```

```
Thr Tyr Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu
65              70              75              80
```

```
Lys His Lys Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser Ser
                85              90              95
```

```
Pro Val Thr Lys Ser Phe Asn Arg Gly Glu Cys
                100             105
```

<210> 111

<211> 642

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized E6 lambda light chain sequence

<400> 111

```
gaaattgtgt tgacacagtc tccagccacc ctgtctttgt ctccagggga aagagccacc      60
ctcacctgca gcgccaccag cagtgttagc tacatccact ggtaccaaca gaggcctggc      120
cagagcccca ggctcctcat ctatagcacc tccaacctgg ccagcgcat cccagccagg      180
```

ttcagtggca gtgggtctgg gagcgactac actctcacca tcagcagcct agagcctgaa	240
gattttgcag tttattactg tcagcagcgt agcagctccc ctttcacctt tggcagcggc	300
accaaagtgg aaattaaagg tcagcccaag gctgccccct cggtcactct gttcccggcc	360
tcctctgagg agcttcaagc caacaaggcc acactggtgt gtctcataag tgacttctac	420
ccgggagccg tgacagtggc ctggaaggca gatagcagcc ccgtcaaggc gggagtggag	480
accaccacac cctccaaaca aagcaacaac aagtacgcgg ccagcagcta tctgagcctg	540
acgcctgagc agtggaaagtc ccacagaagc tacagctgcc aggtcacgca tgaagggagc	600
accgtggaga agacagtggc ccctacagaa tgttcatagt aa	642

<210> 112

<211> 212

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized E6 lambda light chain sequence

<400> 112

Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly	
1                    5                    10                    15	

Glu Arg Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile	
20                    25                    30	

His Trp Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr	
35                    40                    45	

Ser Thr Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser	
50                    55                    60	

Gly Ser Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu	
65                    70                    75                    80	

Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr	
85                    90                    95	

Phe Gly Ser Gly Thr Lys Val Glu Ile Lys Gly Gln Pro Lys Ala Ala	
100                    105                    110	

Pro Ser Val Thr Leu Phe Pro Pro Ser Ser Glu Glu Leu Gln Ala Asn	
115                    120                    125	

Lys Ala Thr Leu Val Cys Leu Ile Ser Asp Phe Tyr Pro Gly Ala Val	
130                    135                    140	

Thr Val Ala Trp Lys Ala Asp Ser Ser Pro Val Lys Ala Gly Val Glu	
145                    150                    155                    160	

Thr Thr Thr Pro Ser Lys Gln Ser Asn Asn Lys Tyr Ala Ala Ser Ser	
165                    170                    175	

Tyr Leu Ser Leu Thr Pro Glu Gln Trp Lys Ser His Arg Ser Tyr Ser  
180 185 190

Cys Gln Val Thr His Glu Gly Ser Thr Val Glu Lys Thr Val Ala Pro  
195 200 205

Thr Glu Cys Ser  
210

<210> 113  
<211> 324  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized lambda light chain constant region sequence

<400> 113  
ggtcagccca aggctgcccc ctcggtcact ctgttccgc cctcctctga ggagcttcaa 60  
gccaacaagg ccacactggt gtgtctcata agtgacttct acccgggagc cgtgacagtg 120  
gcctggaagg cagatagcag ccccgtaag gcgggagtgg agaccaccac accctccaaa 180  
caaagcaaca acaagtacgc ggccagcagc tatctgagcc tgacgcctga gcagtggaag 240  
tcccacagaa gctacagctg ccaggtcacg catgaaggga gcaccgtgga gaagacagtg 300  
gcccctacag aatgttcata gtaa 324

<210> 114  
<211> 106  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized lambda light chain constant region sequence

<400> 114

Gly Gln Pro Lys Ala Ala Pro Ser Val Thr Leu Phe Pro Pro Ser Ser  
1 5 10 15

Glu Glu Leu Gln Ala Asn Lys Ala Thr Leu Val Cys Leu Ile Ser Asp  
20 25 30

Phe Tyr Pro Gly Ala Val Thr Val Ala Trp Lys Ala Asp Ser Ser Pro  
35 40 45

Val Lys Ala Gly Val Glu Thr Thr Thr Pro Ser Lys Gln Ser Asn Asn  
50 55 60

Lys Tyr Ala Ala Ser Ser Tyr Leu Ser Leu Thr Pro Glu Gln Trp Lys  
65 70 75 80

Ser His Arg Ser Tyr Ser Cys Gln Val Thr His Glu Gly Ser Thr Val  
85 90 95

Glu Lys Thr Val Ala Pro Thr Glu Cys Ser  
100 105

<210> 115

<400> 115  
000

<210> 116

<400> 116  
000

<210> 117

<400> 117  
000

<210> 118

<211> 390

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C2 heavy chain variable region sequence

<400> 118

gaggtccagc tggaggagtc agggggaggc ttagtgaagc ctggagggtc cctgaaactc 60

tcctgtgcag cctctggatt cactttcagt ggctatgcca tgtcttgggt tcgccagact 120

ccggagaaga ggctggagtg ggtcgcaacc attagtagtg gtggtactta tatctactat 180

ccagacagtg tgaaggggcg attcaccatc tccagagaca atgccaagaa caccctgtac 240

ctgcaaata gacgtctgag gtctgaggac acggccatgt attactgtgc aagacttggg 300

ggggataatt actacgaata cttcgatgtc tggggcgag ggaccacggt caccgtctcc 360

tccgcaaaa cgacaccccc atctgtctat 390

<210> 119

<211> 130

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C2 heavy chain variable region sequence

<400> 119

Glu Val Gln Leu Glu Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
20 25 30

Ala Met Ser Trp Val Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val  
35 40 45

Ala Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr  
65 70 75 80

Leu Gln Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Met Tyr Tyr Cys  
85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val Trp Gly  
100 105 110

Ala Gly Thr Thr Val Thr Val Ser Ser Ala Lys Thr Thr Pro Pro Ser  
115 120 125

Val Tyr  
130

<210> 120  
<211> 90  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse C2 heavy chain variable framework region 1 (FWR1) sequence

<400> 120  
gaggtccagc tggaggagtc agggggaggc ttagtgaagc ctggagggtc cctgaaactc 60  
tcctgtgcag cctctggatt cactttcagt 90

<210> 121  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse C2 heavy chain variable framework region 1 (FWR1) sequence

<400> 121

Glu Val Gln Leu Glu Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 122  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse C2 heavy chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 122  
ggctatgcc tgtct 15

<210> 123  
<211> 5  
<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C2 heavy chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 123

Gly Tyr Ala Met Ser  
1 5

<210> 124

<211> 42

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C2 heavy chain variable framework region 2 (FWR2) sequence

<400> 124

tgggttcgcc agactccgga gaagaggctg gagtgggtcg ca 42

<210> 125

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C2 heavy chain variable framework region 2 (FWR2) sequence

<400> 125

Trp Val Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val Ala  
1 5 10

<210> 126

<211> 51

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C2 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 126

accattagta gtggtgttac ttatatctac tatccagaca gtgtgaaggg g 51

<210> 127

<211> 17

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C2 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 127

Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys  
1 5 10 15

Gly

<210> 128  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse C2 heavy chain variable framework region 3 (FWR3) sequence  
  
 <400> 128  
 cgattcacca tctccagaga caatgccaaag aacaccctgt acctgcaaat gagcagtcctg 60  
 aggtctgagg acacggccat gtattactgt gcaaga 96  
  
 <210> 129  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse C2 heavy chain variable framework region 3 (FWR3) sequence  
  
 <400> 129  
  
 Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln  
 1 5 10 15  
  
 Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Met Tyr Tyr Cys Ala Arg  
 20 25 30  
  
 <210> 130  
 <211> 36  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse C2 heavy chain variable complementarity determining regions  
 3 (CDR3) sequence  
  
 <400> 130  
 cttggggggg ataattacta cgaatacttc gatgtc 36  
  
 <210> 131  
 <211> 12  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse C2 heavy chain variable complementarity determining regions  
 3 (CDR3) sequence  
  
 <400> 131  
  
 Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val  
 1 5 10  
  
 <210> 132  
  
 <400> 132  
 000  
  
 <210> 133  
  
 <400> 133

000

<210> 134

<400> 134  
000

<210> 135

<400> 135  
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<210> 136

<400> 136  
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<210> 137

<400> 137  
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<210> 138

<400> 138  
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<210> 139

<400> 139  
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<210> 140

<400> 140  
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<210> 141

<400> 141  
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<210> 142

<400> 142  
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<210> 143

<400> 143  
000

<210> 144

<211> 363

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized derived from IGHV3-21\*04 - Humanized C2 heavy chain  
variable region sequence

<400> 144

gaggtgcagc tggaggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60

tcctgtgcag cctctggatt caccttcagt ggctatgcc ttagctgggt ccgccaggct 120

ccaggaagg ggctggagt ggtctcaacc attagtagtg gcggaaccta catatactac 180

cccgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240



```

ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagacttggg      300
ggggataatt actacgaata cttcgatgtc tggggcaaag ggaccacggt caccgtctcc      360
tcc                                          363

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<210>	145
<211>	121
<212>	PRT
<213>	Artificial Sequence

<220>  
<223> Humanized C2 heavy chain variable region sequence

<400> 145

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
20 25 30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val Trp Gly  
100 105 110

Lys Gly Thr Thr Val Thr Val Ser Ser  
115 120

<210>	146
<211>	90
<212>	DNA
<213>	Artificial Sequence

```
<220>
<223> Humanized C2 heavy chain variable framework region 1 (FWR1)
sequence
```

<400> 146  
gaggtgcagc ttgttgagtc tgggggagggc ctggtcaagc ctggggggtc cctgagactc 60

tcctgtgcag cctctggatt caccttcagt 90

<210>	147
<211>	30
<212>	PRT
<213>	Artificial Sequence

<220>  
<223> Humanized C2 heavy chain variable framework region 1 (FWR1)  
sequence

<400> 147

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 148  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 heavy chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 148  
ggctatgcc ttagc 15

<210> 149  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 heavy chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 149

Gly Tyr Ala Met Ser  
1 5

<210> 150  
<211> 43  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 heavy chain variable framework region 2 (FWR2)  
sequence

<400> 150  
tgggtccgcc aggtccagg gaaggggctg gagtgggtct caa 43

<210> 151  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 heavy chain variable framework region 2 (FWR2)  
sequence

<400> 151

Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val Ser  
1 5 10

<210> 152  
 <211> 51  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized C2 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 152  
 accattagta gtggcggaac ctacatatac taccccgact cagtgaaggg c 51

<210> 153  
 <211> 17  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized C2 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 153

Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys  
 1 5 10 15

Gly

<210> 154  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized C2 heavy chain variable framework region 3 (FWR3) sequence  
  
 <400> 154  
 cgattcacca tctccagaga caacgccaaag aactcactgt atctgcaaat gaacagcctg 60  
 agagccgagg acacggccgt gtattactgt gcgaga 96

<210> 155  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized C2 heavy chain variable framework region 3 (FWR3) sequence  
  
 <400> 155

Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr Leu Gln  
 1 5 10 15

Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys Ala Arg  
 20 25 30

<210> 156

<211> 36  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 156  
cttggggggg ataattacta cgaatacttc gatgtc

36

<210> 157  
<211> 12  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 157

Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val  
1 5 10

<210> 158  
<211> 451  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 IgG1 heavy chain sequence

<400> 158

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
20 25 30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val Trp Gly  
100 105 110

Lys Gly Thr Thr Val Thr Val Ser Ser Ala Ser Thr Lys Gly Pro Ser  
115 120 125

Val Phe Pro Leu Ala Pro Ser Ser Lys Ser Thr Ser Gly Gly Thr Ala  
130 135 140

Ala Leu Gly Cys Leu Val Lys Asp Tyr Phe Pro Glu Pro Val Thr Val  
145 150 155 160

Ser Trp Asn Ser Gly Ala Leu Thr Ser Gly Val His Thr Phe Pro Ala  
165 170 175

Val Leu Gln Ser Ser Gly Leu Tyr Ser Leu Ser Ser Val Val Thr Val  
180 185 190

Pro Ser Ser Ser Leu Gly Thr Gln Thr Tyr Ile Cys Asn Val Asn His  
195 200 205

Lys Pro Ser Asn Thr Lys Val Asp Lys Lys Val Glu Pro Lys Ser Cys  
210 215 220

Asp Lys Thr His Thr Cys Pro Pro Cys Pro Ala Pro Glu Leu Leu Gly  
225 230 235 240

Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu Met  
245 250 255

Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser His  
260 265 270

Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val Asp Gly Val Glu Val  
275 280 285

His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn Ser Thr Tyr  
290 295 300

Arg Val Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu Asn Gly  
305 310 315 320

Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Ala Leu Pro Ala Pro Ile  
325 330 335

Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln Val  
340 345 350

Tyr Thr Leu Pro Pro Ser Arg Glu Glu Met Thr Lys Asn Gln Val Ser  
355 360 365

Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val Glu  
370 375 380

Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro Pro  
385 390 395 400

Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr Val

405

410

415

Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met  
 420 425 430

His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu Ser  
 435 440 445

Pro Gly Lys  
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<210> 159

<400> 159  
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<210> 162

<400> 162  
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<210> 163

<211> 1347

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C2 IgG2 heavy chain sequence

<400> 163

gaggtgcagc tgggtggagtc tggggggaggc ctggtcaagc ctgggggggtc cctgagactc 60

tcctgtgcag cctctggatt caccttcagt ggctatgcc a tgagctgggt ccgccaggct 120

ccagggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac 180

cccgaactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240

ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagacttggg 300

ggggataatt actacgaata cttcgatgtc tggggcaaag ggaccacggt caccgtctcc 360

tccgcctcca ccaaggggccc atcggctctc cccctggcgc cctgctccag gagcacctcc 420

gagagcacag ccgccctggg ctgcctggtc aaggactact tccccgaacc ggtgacggtg 480

tcgtggaact caggcgtctt gaccagcggc gtgcacacct tcccagctgt cctacagtcc 540

tcaggactct actccctcag cagcgtgggt accgtgccct ccagcaactt cggcaccag 600

acctacacct gcaacgtaga tcacaagccc agcaacacca aggtggacaa gacagttgag 660

cgcaaatgtt gtgtcgagtg cccaccgtgc ccagcaccac ctgtggcagg accgtcagtc 720

ttcctcttcc ccccaaaacc caaggacacc ctcatgatct cccggacccc tgaggtcacg 780



145		150		155		160
Ser Trp Asn Ser Gly Ala Leu Thr Ser Gly Val His Thr Phe Pro Ala						
	165			170		175
Val Leu Gln Ser Ser Gly Leu Tyr Ser Leu Ser Ser Val Val Thr Val						
	180			185		190
Pro Ser Ser Asn Phe Gly Thr Gln Thr Tyr Thr Cys Asn Val Asp His						
	195			200		205
Lys Pro Ser Asn Thr Lys Val Asp Lys Thr Val Glu Arg Lys Cys Cys						
	210			215		220
Val Glu Cys Pro Pro Cys Pro Ala Pro Pro Val Ala Gly Pro Ser Val						
	225			230		235
Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser Arg Thr						
	245			250		255
Pro Glu Val Thr Cys Val Val Val Asp Val Ser His Glu Asp Pro Glu						
	260			265		270
Val Gln Phe Asn Trp Tyr Val Asp Gly Val Glu Val His Asn Ala Lys						
	275			280		285
Thr Lys Pro Arg Glu Glu Gln Phe Asn Ser Thr Phe Arg Val Val Ser						
	290			295		300
Val Leu Thr Val Val His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys						
	305			310		315
Cys Lys Val Ser Asn Lys Gly Leu Pro Ala Pro Ile Glu Lys Thr Ile						
	325			330		335
Ser Lys Thr Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro						
	340			345		350
Pro Ser Arg Glu Glu Met Thr Lys Asn Gln Val Ser Leu Thr Cys Leu						
	355			360		365
Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn						
	370			375		380
Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro Pro Met Leu Asp Ser						
	385			390		395
Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg						
	405			410		415
Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu						
	420			425		430



His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys  
435 440 445

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<400> 165  
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<210> 166

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<210> 167

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<210> 168  
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<213> Artificial Sequence

<220>  
<223> Mouse C2 light chain variable region sequence

<400> 168  
gacattgtga tcacacagtc tacagcttcc ttaggtgtat ctctggggca gagggccacc 60  
atctcatgca gggccagcaa aagtgtcagt acatctggct atagttatat gcactggtac 120  
caacagagac caggacagcc acccaaactc ctcatctatc ttgcatccaa cctagaatct 180  
ggggtccctg ccaggttcag tggcagtggg tctgggacag acttcaccct caacatccat 240  
cctgtggagg aggaggatgc tgcaacctat tactgtcagc acagtaggga gcttccgttc 300  
acgttcggag gggggaccaa gctggagata aaacgggctg atgctgcacc aactgtatcc 360

<210> 169  
<211> 120  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse C2 light chain variable region sequence

<400> 169

Asp Ile Val Ile Thr Gln Ser Thr Ala Ser Leu Gly Val Ser Leu Gly  
1 5 10 15

Gln Arg Ala Thr Ile Ser Cys Arg Ala Ser Lys Ser Val Ser Thr Ser  
20 25 30

Gly Tyr Ser Tyr Met His Trp Tyr Gln Gln Arg Pro Gly Gln Pro Pro  
35 40 45

Lys Leu Leu Ile Tyr Leu Ala Ser Asn Leu Glu Ser Gly Val Pro Ala  
50 55 60

Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Asn Ile His  
65 70 75 80

Pro Val Glu Glu Glu Asp Ala Ala Thr Tyr Tyr Cys Gln His Ser Arg  
85 90 95

Glu Leu Pro Phe Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys Arg  
100 105 110

Ala Asp Ala Ala Pro Thr Val Ser  
115 120

<210> 170

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C2 light chain variable framework region 1 (FWR1) sequence

<400> 170

gacattgtga tcacacagtc tacagcttcc ttaggtgtat ctctggggca gagggccacc 60

atctcatgc 69

<210> 171

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C2 light chain variable framework region 1 (FWR1) sequence

<400> 171

Asp Ile Val Ile Thr Gln Ser Thr Ala Ser Leu Gly Val Ser Leu Gly  
1 5 10 15

Gln Arg Ala Thr Ile Ser Cys  
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<210> 172

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C2 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 172

agggccagca aaagtgtcag tacatctggc tatagttata tgcac 45

<210> 173

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C2 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 173

Arg	Ala	Ser	Lys	Ser	Val	Ser	Thr	Ser	Gly	Tyr	Ser	Tyr	Met	His
1				5					10					15

<210> 174

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C2 light chain variable framework region 2 (FWR2) sequence

<400> 174

tggtaccaac agagaccagg acagccaccc aaactcctca tctat 45

<210> 175

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C2 light chain variable framework region 2 (FWR2) sequence

<400> 175

Trp	Tyr	Gln	Gln	Arg	Pro	Gly	Gln	Pro	Pro	Lys	Leu	Leu	Ile	Tyr
1				5					10					15

<210> 176

<211> 20

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C2 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 176

cttgcattcca acctagaatc 20

<210> 177

<211> 7

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C2 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 177

Leu	Ala	Ser	Asn	Leu	Glu	Ser
1				5		

<210> 178

<211> 97

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C2 light chain variable framework region 3 (FWR3) sequence

<400> 178  
 tggggtcct gccaggttca gtggcagtgg gtctgggaca gatttcaccc tcaacatcca 60  
 tcctgtggag gaggaggatg ctgcaaccta ttactgt 97

<210> 179  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse C2 light chain variable framework region 3 (FWR3) sequence

<400> 179  
 Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 1 5 10 15  
 Leu Asn Ile His Pro Val Glu Glu Glu Asp Ala Ala Thr Tyr Tyr Cys  
 20 25 30

<210> 180  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse C2 light chain variable complementarity determining regions  
 3 (CDR3) sequence

<400> 180  
 cagcacagta gggagcttcc gttcacg 27

<210> 181  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse C2 light chain variable complementarity determining regions  
 3 (CDR3) sequence

<400> 181  
 Gln His Ser Arg Glu Leu Pro Phe Thr  
 1 5

<210> 182  
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<400> 193  
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<210> 194

<211> 339

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized derived from IGKV73\*01 - Humanized C2 light chain  
variable region sequence

<400> 194  
gacattgtgc tgacccagtc tccagcctcc ttggccgtgt ctccaggaca gagggccacc 60  
atcacctgca gagccagtaa gagtgtcagt accagcggat actcctacat gactggtat 120  
cagcagaaac caggacaacc tcctaaactc ctgatttacc tggcatccaa tctggagagc 180  
gggggtcccag ccaggttcag cggcagtggg tctgggaccg atttcaccct cacaattaat 240  
cctgtggaag ctaatgatac tgcaaattat tactgtcagc acagtaggga gctgcctttc 300  
acattcggcg gagggaccaa ggtggagatc aaacgaact 339

<210> 195

<211> 113

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized derived from IGKV73\*01 - Humanized C2 light chain variable region sequence

<400> 195

Asp Ile Val Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly  
1 5 10 15

Gln Arg Ala Thr Ile Thr Cys Arg Ala Ser Lys Ser Val Ser Thr Ser  
20 25 30

Gly Tyr Ser Tyr Met His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro  
35 40 45

Lys Leu Leu Ile Tyr Leu Ala Ser Asn Leu Glu Ser Gly Val Pro Ala  
50 55 60

Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn  
65 70 75 80

Pro Val Glu Ala Asn Asp Thr Ala Asn Tyr Tyr Cys Gln His Ser Arg  
85 90 95

Glu Leu Pro Phe Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg  
100 105 110

Thr

<210> 196

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C2 light chain variable framework region 1 (FWR1) acid sequence

<400> 196

gacattgtgc tgaccagtc tccagcctcc ttggccgtgt ctccaggaca gagggccacc 60

atcacctgc 69

<210> 197

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C2 light chain variable framework region 1 (FWR1) acid sequence

<400> 197

Asp Ile Val Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly  
1 5 10 15

Gln Arg Ala Thr Ile Thr Cys  
20

<210> 198  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 198  
agagccagta agagtgtcag taccagcgga tactcctaca tgcac 45

<210> 199  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 199

Arg Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser Tyr Met His  
1 5 10 15

<210> 200  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 heavy light variable framework region 2 (FWR2) acid  
sequence

<400> 200  
tggatcagc agaaaccagg acaacctcct aaactcctga ttac 45

<210> 201  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 heavy light variable framework region 2 (FWR2) acid  
sequence

<400> 201

Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 202  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable complementarity determining  
regions 2 (CDR2) sequence

<400> 202  
ctggcatcca atctggagag c 21

<210> 203  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable complementarity determining  
regions 2 (CDR2) sequence

<400> 203

Leu Ala Ser Asn Leu Glu Ser  
1 5

<210> 204  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable framework region 3 (FWR3) acid  
sequence

<400> 204  
ggggtcccag ccaggttcag cggcagtggg tctgggaccg atttcaccct cacaattaat 60

cctgtggaag ctaatgatac tgcaaattat tactgt 96

<210> 205  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable framework region 3 (FWR3) acid  
sequence

<400> 205

Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Thr Ile Asn Pro Val Glu Ala Asn Asp Thr Ala Asn Tyr Tyr Cys  
20 25 30

<210> 206  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable complementarity determining  
regions 3 (CDR3) sequence

<400> 206  
cagcacagta gggagctgcc tttcaca 27

<210> 207  
<211> 9



<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 207

Gln His Ser Arg Glu Leu Pro Phe Thr  
1 5

<210> 208  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 208  
ctgcagagta agaattttcc tcccaca

27

<210> 209  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 209

Leu Gln Ser Lys Asn Phe Pro Pro Thr  
1 5

<210> 210

<400> 210  
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<210> 211

<400> 211  
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<210> 212

<400> 212  
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<400> 220  
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<210> 221

<400> 221  
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<210> 222  
<211> 63  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Murine Ig kappa chain leader sequence

<400> 222  
atggagacag acacactcct gctatgggta ctgctgctct gggttccagg ttccactggt 60  
gac 63

<210> 223  
<211> 21  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Murine Ig kappa chain leader sequence

<400> 223

Met Glu Thr Asp Thr Leu Leu Leu Trp Val Leu Leu Leu Trp Val Pro  
1 5 10 15

Gly Ser Thr Gly Asp  
20

<210> 224  
<211> 60  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Interleukin2 (IL2) leader sequence  
  
<400> 224  
atgtacagga tgcaactcct gtcttgcatt gcactaagtc ttgcacttgt cacaaacagt 60

<210> 225  
<211> 20  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Interleukin2 (IL2) leader sequence  
  
<400> 225

Met Tyr Arg Met Gln Leu Leu Ser Cys Ile Ala Leu Ser Leu Ala Leu  
1 5 10 15

Val Thr Asn Ser  
20

<210> 226  
<211> 48  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD33 leader sequence

<400> 226  
atgcctcttc tgcttctgct tcctctgctt tgggctggag ctcttgct 48

<210> 227  
<211> 16  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CD33 leader sequence

<400> 227

Met Pro Leu Leu Leu Leu Pro Leu Leu Trp Ala Gly Ala Leu Ala  
1 5 10 15

<210> 228  
<211> 57  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> IGHV321\*03 leader sequence

<400> 228  
atggaactgg ggctccgctg ggttttcctt gttgctatatt tagaagggtg ccagtggt 57

<210> 229  
<211> 19  
<212> PRT  
<213> Artificial Sequence

<220>

<223> IGHV321\*03 leader sequence

<400> 229

Met Glu Leu Gly Leu Arg Trp Val Phe Leu Val Ala Ile Leu Glu Gly  
1 5 10 15

Val Gln Cys

<210> 230

<211> 60

<212> DNA

<213> Artificial Sequence

<220>

<223> IGHV311\*02 leader sequence

<400> 230

atggaagccc cagcgcagct tctcttcctc ctgctactct ggctcccaga taccactgga 60

<210> 231

<211> 20

<212> PRT

<213> Artificial Sequence

<220>

<223> IGHV311\*02 leader sequence

<400> 231

Met Glu Ala Pro Ala Gln Leu Leu Phe Leu Leu Leu Leu Trp Leu Pro  
1 5 10 15

Asp Thr Thr Gly  
20

<210> 232

<211> 729

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized E6 single chain GS3

<400> 232

gaggtgcagc tgggtggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60

tcctgtgcag cctctggatt cacccttcagt aggtatggca tgagctgggt ccgccaggct 120

ccagggaaga ggctggagtg ggtctcaacc attagtggcg gaggcaccta catatactac 180

ccagactcag tgaagggccg attcaccatc tccagagaca acgccaagaa caccctgtat 240

ctgcaaatac acagcctgag agccgaggac acggcttgtt attactgtac cagagataac 300

tatggccgca actatgatta tggcatggat tattggggcc agggcaccct ggtgaccgtg 360

agcagcggcg gtggcggatc cggcgggtggc ggatccggcg gtggcggatc cgaaattgtg 420

ttgacacagt ctccagccac cctgtctttg tctccagggg aaagagccac cctcacctgc 480

agcgccacca gcagtgttag ctacatccac tggtagcaac agaggcctgg ccagagcccc 540

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aggctcctca tctatagcac ctccaacctg gccagcggca tcccagccag gttcagtggc      600
agtgggtctg ggagcgacta cactctcacc atcagcagcc tagagcctga agattttgca      660
gtttattact gtcagcagcg tagcagctcc cttttcacct ttggcagcgg caccaaagtg      720
gaaattaaa                                     729

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Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys Arg Leu Glu Trp Val  
35 40 45

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr  
65 70 75 80

Thr Arg Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr Gly Met Asp Tyr Trp  
100 105 110

Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Ile Val Leu Thr Gln Ser  
130 135 140

Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp Tyr Gln Gln Arg Pro  
165 170 175

Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Ser Asp Tyr Thr  
195 200 205

Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys  
210 215 220

Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly Ser Gly Thr Lys Val  
225 230 235 240

Glu Ile Lys

<210> 234  
<211> 747  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized E6 single chain IgG1noC

<400> 234  
gaggtgcagc tgggtggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60  
tcctgtgcag cctctggatt caccttcagt aggtatggca tgagctgggt ccgccaggct 120  
ccagggaaga ggctggagtg ggtctcaacc attagtggcg gaggcaccta catatactac 180  
ccagactcag tgaagggccg attcaccatc tccagagaca acgccaagaa caccctgtat 240  
ctgcaaatac acagcctgag agccgaggac acggctgtgt attactgtac cagagataac 300  
tatggccgca actatgatta tggcatggat tattggggcc agggcaccct ggtgaccgtg 360  
agcagcgata aaaccatac taaaccgcca aaaccggcgc cggaactgct ggggtggtcct 420  
ggtaccggtg aaattgtgtt gacacagtct ccagccaccc tgtctttgtc tccaggggaa 480  
agagccaccc tcacctgcag cgccaccagc agtgtttagct acatccactg gtaccaacag 540  
aggcctggcc agagccccag gtcctcatc tatagcacct ccaacctggc cagcggcatc 600  
ccagccaggc tcagtggcag tgggtctggg agcgactaca ctctcaccat cagcagccta 660  
gagcctgaag attttgcagt ttattactgt cagcagcgta gcagctcccc tttcaccttt 720  
ggcagcggca ccaaagtga aattaa 747

<210> 235  
<211> 249  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized E6 single chain IgG1noC

<400> 235

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Arg Tyr  
20 25 30

Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys Arg Leu Glu Trp Val

35

40

45

Ser Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr  
65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Thr Arg Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr Gly Met Asp Tyr Trp  
100 105 110

Gly Gln Gly Thr Leu Val Thr Val Ser Ser Asp Lys Thr His Thr Lys  
115 120 125

Pro Pro Lys Pro Ala Pro Glu Leu Leu Gly Gly Pro Gly Thr Gly Glu  
130 135 140

Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly Glu  
145 150 155 160

Arg Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile His  
165 170 175

Trp Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr Ser  
180 185 190

Thr Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly  
195 200 205

Ser Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp  
210 215 220

Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe  
225 230 235 240

Gly Ser Gly Thr Lys Val Glu Ile Lys  
245

<210> 236

<211> 813

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized E6 single chain X4 (linker is IgG1 and IgG2 modified hinge region)

<400> 236

gaggtgcagc tgggtggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60

tcctgtgcag cctctggatt caccttcagt aggtatggca tgagctgggt ccgccaggct 120





130

135

140

Gly Gly Pro Thr Ile Lys Pro Pro Lys Pro Pro Lys Pro Ala Pro Asn  
145 150 155 160

Leu Leu Gly Gly Pro Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu  
165 170 175

Ser Leu Ser Pro Gly Glu Arg Ala Thr Leu Thr Cys Ser Ala Thr Ser  
180 185 190

Ser Val Ser Tyr Ile His Trp Tyr Gln Gln Arg Pro Gly Gln Ser Pro  
195 200 205

Arg Leu Leu Ile Tyr Ser Thr Ser Asn Leu Ala Ser Gly Ile Pro Ala  
210 215 220

Arg Phe Ser Gly Ser Gly Ser Gly Ser Asp Tyr Thr Leu Thr Ile Ser  
225 230 235 240

Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser  
245 250 255

Ser Ser Pro Phe Thr Phe Gly Ser Gly Thr Lys Val Glu Ile Lys  
260 265 270

<210> 238  
<211> 747  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 single chain GS3

<400> 238  
gaggtgcagc tgggtggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60  
tcctgtgcag cctctggatt caccttcagt ggctatgcc a tgagctgggt ccgccaggct 120  
ccagggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac 180  
cccgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240  
ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagacttggg 300  
ggggataatt actacgaata cttcgatgtc tggggcaaag ggaccacggt caccgtctcc 360  
tccggcgggtg gcggatccgg cgggtggcgga tccggcgggtg gcggatccga cattgtgctg 420  
accagttctc cagcctcctt ggccgtgtct ccaggacaga gggccacat cacctgcaga 480  
gccagtaaga gtgtcagtac cagcggatac tcctacatgc actggtatca gcagaaacca 540  
ggacaacctc ctaaactcct gatttacctg gcatccaatc tggagagcgg ggtcccagcc 600  
aggttcagcg gcagtgggtc tgggaccgat ttcaccctca caattaatcc tgtggaagct 660  
aatgatactg caaattatta ctgtcagcac agtagggagc tgcctttcac attcggcgga 720  
gggaccaagg tggagatcaa acgaact 747

<210> 239  
<211> 249  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 single chain GS3

<400> 239

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
20 25 30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val Trp Gly  
100 105 110

Lys Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly  
115 120 125

Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Leu Thr Gln Ser Pro  
130 135 140

Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala Thr Ile Thr Cys Arg  
145 150 155 160

Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser Tyr Met His Trp Tyr  
165 170 175

Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Leu Ala Ser  
180 185 190

Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly  
195 200 205

Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu Ala Asn Asp Thr Ala  
210 215 220

Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro Phe Thr Phe Gly Gly

225 230 235 240

Gly Thr Lys Val Glu Ile Lys Arg Thr  
245

<210> 240

<400> 240  
000

<210> 241

<400> 241  
000

<210> 242  
<211> 831  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C2 single chain X4 (linker is IgG1 and IgG2 modified  
hinge region

<400> 242  
gaggtgcagc tgggtggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60  
tcctgtgcag cctctggatt caccttcagt ggctatgcc a tgagctgggt ccgccaggct 120  
ccagggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac 180  
cccgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240  
ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagacttggg 300  
ggggataatt actacgaata cttcgatgtc tggggcaaa ggaccacggt caccgtctcc 360  
tccgataaaa cccatactaa accgcaaaa ccggcgccgg aactgctggg tggtcctggt 420  
accggtactg gtggtccgac tattaacct ccgaaacctc cgaaacctgc tccgaacctg 480  
ctgggtggtc cggacattgt gctgaccag tctccagcct ccttggccgt gtctccagga 540  
cagagggccca ccatcacctg cagagccagt aagagtgtca gtaccagcgg atactcctac 600  
atgcactggt atcagcagaa accaggacaa cctcctaaac tcctgattta cctggcatcc 660  
aatctggaga gcgggggtccc agccaggttc agcggcagtg ggtctgggac cgatttcacc 720  
ctcacaatta atcctgtgga agctaattgat actgcaaatt attactgtca gcacagtagg 780  
gagctgcctt tcacattcgg cggaggggacc aaggtggaga tcaaacgaac t 831

<210> 243  
<211> 277  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C2 single chain X4 (linker is IgG1 and IgG2 modified  
hinge region

<400> 243

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
 20 25 30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
 35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
 50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
 65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
 85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val Trp Gly  
 100 105 110

Lys Gly Thr Thr Val Thr Val Ser Ser Asp Lys Thr His Thr Lys Pro  
 115 120 125

Pro Lys Pro Ala Pro Glu Leu Leu Gly Gly Pro Gly Thr Gly Thr Gly  
 130 135 140

Gly Pro Thr Ile Lys Pro Pro Lys Pro Pro Lys Pro Ala Pro Asn Leu  
 145 150 155 160

Leu Gly Gly Pro Asp Ile Val Leu Thr Gln Ser Pro Ala Ser Leu Ala  
 165 170 175

Val Ser Pro Gly Gln Arg Ala Thr Ile Thr Cys Arg Ala Ser Lys Ser  
 180 185 190

Val Ser Thr Ser Gly Tyr Ser Tyr Met His Trp Tyr Gln Gln Lys Pro  
 195 200 205

Gly Gln Pro Pro Lys Leu Leu Ile Tyr Leu Ala Ser Asn Leu Glu Ser  
 210 215 220

Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 225 230 235 240

Leu Thr Ile Asn Pro Val Glu Ala Asn Asp Thr Ala Asn Tyr Tyr Cys  
 245 250 255

Gln His Ser Arg Glu Leu Pro Phe Thr Phe Gly Gly Gly Thr Lys Val  
 260 265 270

Glu Ile Lys Arg Thr  
 275

<210> 244  
<211> 744  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 single chain GS3

<400> 244  
caggttcagc tggatgcagtc tggagctgag gtgaagaagc ctggggcctc agtgaaggtc 60  
tcctgcaagg cttctggtta cacctttacc gactacgcc tgaactgggt gcgacaggcc 120  
cctggacaag ggcttgagt gatgggagt atcagcacct tcagcggtaa cacaacttc 180  
aaccagaagt tcaagggcag agtcaccatg accacagaca catccacgag cacagcctac 240  
atggagctga ggagcctgag atctgacgac acggccgtgt attactgtgc gagaagcgac 300  
tactacggcc catacttcga ctactggggc cagggcacca ccctgaccgt gtccagcggc 360  
ggatggcggt ccggcggtgg cggatccggc ggtggcggt ccgatattgt gatgaccag 420  
actccactct ctctgtccgt cacccttga cagccggcct ccatctcctg caggtctagt 480  
cagaccattg tccatagtaa tggaaacacc tatttgaggt ggtacctgca gaagccaggc 540  
cagtctccac agctcctgat ctataaggtt tccaaccggt tctctggagt gccagatagg 600  
ttcagtggca gcgggtcagg gacagatttc aactgaaaa tcagccgggt ggaggctgag 660  
gatgttgggg tttattactg cttccaaggt agccacgtgc ctttcacctt cggcggaggg 720  
accaaggtgg agatcaaacg aact 744

<210> 245  
<211> 248  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 single chain GS3

<400> 245

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ala  
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asp Tyr  
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met  
35 40 45

Gly Val Ile Ser Thr Phe Ser Gly Asn Thr Asn Phe Asn Gln Lys Phe  
50 55 60

Lys Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr  
65 70 75 80

Met Glu Leu Arg Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Ala Arg Ser Asp Tyr Tyr Gly Pro Tyr Phe Asp Tyr Trp Gly Gln Gly  
100 105 110

Thr Thr Leu Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly  
115 120 125

Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Thr Pro Leu Ser  
130 135 140

Leu Ser Val Thr Pro Gly Gln Pro Ala Ser Ile Ser Cys Arg Ser Ser  
145 150 155 160

Gln Thr Ile Val His Ser Asn Gly Asn Thr Tyr Leu Glu Trp Tyr Leu  
165 170 175

Gln Lys Pro Gly Gln Ser Pro Gln Leu Leu Ile Tyr Lys Val Ser Asn  
180 185 190

Arg Phe Ser Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr  
195 200 205

Asp Phe Thr Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Val Gly Val  
210 215 220

Tyr Tyr Cys Phe Gln Gly Ser His Val Pro Phe Thr Phe Gly Gly Gly  
225 230 235 240

Thr Lys Val Glu Ile Lys Arg Thr  
245

<210> 246

<400> 246  
000

<210> 247

<400> 247  
000

<210> 248

<211> 828

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C3 single chain X4 (linker is IgG1 and IgG2 modified  
hinge region

<400> 248

caggttcagc tgggtgcagtc tggagctgag gtgaagaagc ctggggcctc agtgaaggtc 60

tcctgcaagg cttctggtta cacctttacc gactacgcca tgaactgggt gcgacaggcc 120

cctggacaag ggcttgagt gatgggagt atcagcacct tcagcggtaa cacaaacttc 180

aaccagaagt tcaagggcag agtcaccatg accacagaca catccacgag cacagcctac 240

```

atggagctga ggagcctgag atctgacgac acggccgtgt attactgtgc gagaagcgac      300
tactacggcc catacttcga ctactggggc cagggcacca ccctgaccgt gtccagcgat      360
aaaaccata ctaaaccgcc aaaaccggcg ccggaactgc tgggtggtcc tggtagcggt      420
actggtggtc cgactattaa acctccgaaa cctccgaaac ctgctccgaa cctgctgggt      480
ggtccggata ttgtgatgac ccagactcca ctctctctgt ccgtcacccc tggacagccg      540
gcctccatct cctgcaggtc tagtcagacc attgtccata gtaatggaaa cacctatttg      600
gagtgggtacc tgcagaagcc aggccagtct ccacagctcc tgatctataa ggtttccaac      660
cggttctctg gagtgccaga taggttcagt ggcagcgggt cagggacaga tttcacactg      720
aaaatcagcc gggtaggggc tgaggatgtt ggggtttatt actgcttcca aggtagccac      780
gtgcctttca ctttcggcgg agggaccaag gtggagatca aacgaact      828

```

<210> 249

<211> 276

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C3 single chain X4 (linker is IgG1 and IgG2 modified hinge region)

<400> 249

```

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ala
1              5              10              15

```

```

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asp Tyr
          20              25              30

```

```

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
          35              40              45

```

```

Gly Val Ile Ser Thr Phe Ser Gly Asn Thr Asn Phe Asn Gln Lys Phe
50              55              60

```

```

Lys Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
65              70              75              80

```

```

Met Glu Leu Arg Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
          85              90              95

```

```

Ala Arg Ser Asp Tyr Tyr Gly Pro Tyr Phe Asp Tyr Trp Gly Gln Gly
          100              105              110

```

```

Thr Thr Leu Thr Val Ser Ser Asp Lys Thr His Thr Lys Pro Pro Lys
          115              120              125

```

```

Pro Ala Pro Glu Leu Leu Gly Gly Pro Gly Thr Gly Thr Gly Gly Pro
          130              135              140

```

Thr Ile Lys Pro Pro Lys Pro Pro Lys Pro Ala Pro Asn Leu Leu Gly  
145 150 155 160

Gly Pro Asp Ile Val Met Thr Gln Thr Pro Leu Ser Leu Ser Val Thr  
165 170 175

Pro Gly Gln Pro Ala Ser Ile Ser Cys Arg Ser Ser Gln Thr Ile Val  
180 185 190

His Ser Asn Gly Asn Thr Tyr Leu Glu Trp Tyr Leu Gln Lys Pro Gly  
195 200 205

Gln Ser Pro Gln Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser Gly  
210 215 220

Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu  
225 230 235 240

Lys Ile Ser Arg Val Glu Ala Glu Asp Val Gly Val Tyr Tyr Cys Phe  
245 250 255

Gln Gly Ser His Val Pro Phe Thr Phe Gly Gly Gly Thr Lys Val Glu  
260 265 270

Ile Lys Arg Thr  
275

<210> 250  
<211> 741  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C8 single chain GS3 (linker is [Gly4Ser1]3

<400> 250  
gaggtgcagc tgggtggagtc tgggggaggc ctggtcaagc ctggggggtc cctgagactc 60  
tcctgtgcag cctctggatt caccttcagt ggctatgcc a tgagctgggt ccgccaggct 120  
ccagggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac 180  
cctgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240  
ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagactgggc 300  
ggcgataact attatgaata ttggggcaaa gggaccacgg tcaccgtctc ctccggcggt 360  
ggcggatccg gcggtggcgg atccggcggt ggcggatccg acatcgtgat gaccagctct 420  
ccagactccc tggctgtgtc tctgggcgag agggccacca tcaactgcag ggccagcaag 480  
agtgttagca ccagcggcta cagctacatg cactggtacc agcagaaacc aggacagcct 540  
cctaagctgc tcatttacct ggtgtctaac ctggaatccg gggtcctga ccgattcagt 600  
ggcagcgggt ctgggacaga ttctactctc accatcagca gcctgcaggc tgaagatgtg 660  
gcagtttatt actgtcaaca cattcgggaa ctgaccagga gtgaattcgg cggagggacc 720



&lt;210&gt; 251

&lt;211&gt; 247

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Humanized C8 single chain GS3 (linker is [Gly4Ser1]3

&lt;400&gt; 251

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
20 25 30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Trp Gly Lys Gly Thr  
100 105 110

Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser  
115 120 125

Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu  
130 135 140

Ala Val Ser Leu Gly Glu Arg Ala Thr Ile Asn Cys Arg Ala Ser Lys  
145 150 155 160

Ser Val Ser Thr Ser Gly Tyr Ser Tyr Met His Trp Tyr Gln Gln Lys  
165 170 175

Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Leu Val Ser Asn Leu Glu  
180 185 190

Ser Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe  
195 200 205

Thr Leu Thr Ile Ser Ser Leu Gln Ala Glu Asp Val Ala Val Tyr Tyr  
210 215 220

Cys Gln His Ile Arg Glu Leu Thr Arg Ser Glu Phe Gly Gly Gly Thr  
225 230 235 240

Lys Val Glu Ile Lys Arg Thr  
245

<210> 252

<400> 252  
000

<210> 253

<400> 253  
000

<210> 254

<400> 254  
000

<210> 255

<400> 255  
000

<210> 256

<211> 1509

<212> DNA

<213> Artificial Sequence

<220>

<223> pSECTag2 E6 scFVFC

<400> 256

atggagacag acacactcct gctatgggta ctgctgctct gggttccagg ttccactggt 60

gacgcggccc agccggccga ggtgcagctg gtggagtctg ggggaggcct ggtcaagcct 120

gggggggtccc tgagactctc ctgtgcagcc tctggattca ccttcagtag gtatggcatg 180

agctgggtcc gccaggctcc agggaagagg ctggagtggg tctcaaccat tagtggcgga 240

ggcacctaca tatactaccc agactcagtg aagggccgat tcaccatctc cagagacaac 300

gccaagaaca ccctgtatct gcaaatgaac agcctgagag ccgaggacac ggctgtgtat 360

tactgtacca gagataacta tggccgcaac tatgattatg gcatggatta ttggggccag 420

ggcaccttgg tgaccgtgag cagcggcggt ggccgatccg gcggtggcgg atccggcggt 480

ggcggatccg aaattgtgtt gacacagtct ccagccaccc tgtctttgtc tccaggggaa 540

agagccaccc tcacctgcag cgccaccagc agtgtttagct acatccactg gtaccaacag 600

aggcctggcc agagccccag gtcctcatc tatagcacct ccaacctggc cagcggcatc 660

ccagccaggt tcagtggcag tgggtctggg agcgactaca ctctcaccat cagcagccta 720

gagcctgaag attttgcagt ttattactgt cagcagcgta gcagctcccc tttcaccttt 780

ggcagcggca ccaaagtgga aattaaagag cccaaatctt gtgacaaaac tcacacatgc 840

ccaccgtgcc cagcacctga actcctgggg ggaccgtcag tcttcctctt cccccaaaa 900

cccaaggaca cctcatgat ctcccgacc cctgaggtca catgcgtggt ggtggacgtg 960

agccacgaag accctgaggt caagttcaac tggtagctgg acggcgtgga ggtgcataat 1020  
gccaagacaa agccgcggga ggagcagtag aacagcacgt accgtgtggt cagcgtcctc 1080  
accgtcctgc accaggactg gctgaatggc aaggagtaca agtgcaaggt ctccaacaaa 1140  
gccctcccag ccccatcgga gaaaaccatc tccaaagcca aagggcagcc ccgagaacca 1200  
caggtgtaca ccctgcccc atcccgggag gagatgacca agaaccaggt cagcctgacc 1260  
tgcctgggtca aaggcttcta tcccagcgac atcgccgtgg agtgggagag caatgggcag 1320  
ccggagaaca actacaagac cacgcctccc gtgctggact ccgacggctc cttcttcctc 1380  
tacagcaagc tcaccgtgga caagagcagg tggcagcagg ggaacgtctt ctcatgctcc 1440  
gtgatgcatg aggctctgca caaccactac acgcagaaga gcctctccct gtctccgggt 1500  
aaatgataa 1509

<210> 257  
<211> 501  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> pSECTag2 E6 scFVFC

<400> 257

Met Glu Thr Asp Thr Leu Leu Leu Trp Val Leu Leu Leu Trp Val Pro  
1 5 10 15  
  
Gly Ser Thr Gly Asp Ala Ala Gln Pro Ala Glu Val Gln Leu Val Glu  
20 25 30  
  
Ser Gly Gly Gly Leu Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys  
35 40 45  
  
Ala Ala Ser Gly Phe Thr Phe Ser Arg Tyr Gly Met Ser Trp Val Arg  
50 55 60  
  
Gln Ala Pro Gly Lys Arg Leu Glu Trp Val Ser Thr Ile Ser Gly Gly  
65 70 75 80  
  
Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile  
85 90 95  
  
Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln Met Asn Ser Leu  
100 105 110  
  
Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys Thr Arg Asp Asn Tyr Gly  
115 120 125  
  
Arg Asn Tyr Asp Tyr Gly Met Asp Tyr Trp Gly Gln Gly Thr Leu Val  
130 135 140  
  
Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly  
145 150 155 160

Gly Gly Ser Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu  
 165 170 175

Ser Pro Gly Glu Arg Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val  
 180 185 190

Ser Tyr Ile His Trp Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu  
 195 200 205

Leu Ile Tyr Ser Thr Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe  
 210 215 220

Ser Gly Ser Gly Ser Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu  
 225 230 235 240

Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser  
 245 250 255

Pro Phe Thr Phe Gly Ser Gly Thr Lys Val Glu Ile Lys Glu Pro Lys  
 260 265 270

Ser Cys Asp Lys Thr His Thr Cys Pro Pro Cys Pro Ala Pro Glu Leu  
 275 280 285

Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr  
 290 295 300

Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val  
 305 310 315 320

Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val Asp Gly Val  
 325 330 335

Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn Ser  
 340 345 350

Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu  
 355 360 365

Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Ala Leu Pro Ala  
 370 375 380

Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro  
 385 390 395 400

Gln Val Tyr Thr Leu Pro Pro Ser Arg Glu Glu Met Thr Lys Asn Gln  
 405 410 415

Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala  
 420 425 430

Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr  
435 440 445

Pro Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu  
450 455 460

Thr Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser  
465 470 475 480

Val Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser  
485 490 495

Leu Ser Pro Gly Lys  
500

<210> 258

<400> 258  
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<210> 259

<400> 259  
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<210> 260

<400> 260  
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<210> 261

<400> 261  
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<210> 262

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<210> 270

<400> 270  
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<210> 271

<400> 271  
000

<210> 272  
<211> 702  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Human IgG1 Fc sequence

<400> 272  
gagcccaaatt cttgtgacaa aactcacaca tgcccaccgt gccagcacc tgaactcctg 60  
gggggaccgt cagtcttcct cttcccccca aaaccaagg acaccctcat gatctcccgg 120  
acccttgagg tcacatgcgt ggtggtggac gtgagccacg aagaccctga ggtcaagttc 180  
aactggtacg tggacggcgt ggagggtgcat aatgccaaga caaagccgcg ggaggagcag 240  
tacaacagca cgtaccgtgt ggtcagcgtc ctcaccgtcc tgcaccagga ctggctgaat 300  
ggcaaggagt acaagtgcaa ggtctccaac aaagccctcc cagcccccat cgagaaaacc 360  
atctccaaag ccaaagggca gcccagagaa ccacaggtgt acaccctgcc cccatcccgg 420  
gaggagatga ccaagaacca ggtcagcctg acctgcctgg tcaaaggctt ctatcccagc 480  
gacatcgccg tggagtggga gagcaatggg cagccggaga acaactacaa gaccacgcct 540  
cccgtgctgg actccgacgg ctccttcttc ctctacagca agctcaccgt ggacaagagc 600  
aggtggcagc aggggaacgt cttctcatgc tccgtgatgc atgaggctct gcacaaccac 660  
tacacgcaga agagcctctc cctgtctccg ggtaaattgat aa 702

<210> 273  
<211> 232  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Human IgG1 Fc sequence

<400> 273

Glu Pro Lys Ser Cys Asp Lys Thr His Thr Cys Pro Pro Cys Pro Ala  
1 5 10 15

Pro Glu Leu Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro  
 20 25 30  
 Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val  
 35 40 45  
 Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val  
 50 55 60  
 Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln  
 65 70 75 80  
 Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln  
 85 90 95  
 Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Ala  
 100 105 110  
 Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro  
 115 120 125  
 Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Arg Glu Glu Met Thr  
 130 135 140  
 Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser  
 145 150 155 160  
 Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr  
 165 170 175  
 Lys Thr Thr Pro Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr  
 180 185 190  
 Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe  
 195 200 205  
 Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys  
 210 215 220  
 Ser Leu Ser Leu Ser Pro Gly Lys  
 225 230

<210> 274  
 <211> 666  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Human IgG1 CH2CH3 domain sequence

<400> 274  
 ccgtgccag cacctgaact cctgggggga ccgtcagtct tcctcttccc cccaaaaccc 60  
 aaggacaccc tcatgatctc ccggaccct gaggtcacat gcgtggtggt ggacgtgagc 120

cacgaagacc ctgagggtcaa gttcaactgg tacgtggacg gcgtggaggt gcataatgcc	180
aagacaaagc cgcgggagga gcagtacaac agcacgtacc gtgtgggtcag cgtcctcacc	240
gtcctgcacc aggactggct gaatggcaag gagtacaagt gcaagggtctc caacaaagcc	300
ctcccagccc ccatcgagaa aaccatctcc aaagccaaag ggcagccccg agaaccacag	360
gtgtacaccc tgcccccatc ccgggaggag atgaccaaga accagggtcag cctgacctgc	420
ctgggtcaaag gcttctatcc cagcgacatc gccgtggagt gggagagcaa tgggcagccg	480
gagaacaact acaagaccac gcctcccgtg ctggactccg acggctcctt cttcctctac	540
agcaagctca ccgtggacaa gagcagggtgg cagcagggga acgtcttctc atgctccgtg	600
atgcatgagg ctctgcacaa ccactacacg cagaagagcc tctccctgtc tccgggtaaa	660
tgataa	666

<210> 275

<211> 220

<212> PRT

<213> Artificial Sequence

<220>

<223> Human IgG1 CH2CH3 domain sequence

<400> 275

Pro Cys Pro Ala Pro Glu Leu Leu Gly Gly Pro Ser Val Phe Leu Phe	
1 5 10 15	

Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val	
20 25 30	

Thr Cys Val Val Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe	
35 40 45	

Asn Trp Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro	
50 55 60	

Arg Glu Glu Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr	
65 70 75 80	

Val Leu His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val	
85 90 95	

Ser Asn Lys Ala Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Ala	
100 105 110	

Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Arg	
115 120 125	

Glu Glu Met Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly	
130 135 140	

Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro



145	150	155	160
Glu Asn Asn Tyr Lys Thr Thr Pro Pro Val Leu Asp Ser Asp Gly Ser	165	170	175
Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln Gln	180	185	190
Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His	195	200	205
Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys	210	215	220

<210> 276  
 <211> 327  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Human IgG1 CH3 domain sequence

<400> 276	
gggcagcccc gagaaccaca ggtgtacacc ctgcccccat cccgggagga gatgaccaag	60
aaccagggtca gcctgacctg cctggtcaaa ggcttctatc ccagcgacat cgccgtggag	120
tgggagagca atgggcagcc ggagaacaac tacaagacca cgcctccgt gctggactcc	180
gacggctcct tcttcctcta cagcaagctc accgtggaca agagcagggtg gcagcagggg	240
aacgtcttct catgctccgt gatgcatgag gctctgcaca accactacac gcagaagagc	300
ctctccctgt ctccgggtaa atgataa	327

<210> 277  
 <211> 107  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Human IgG1 CH3 domain sequence

<400> 277
Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Arg Glu
1 5 10 15
Glu Met Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly Phe
20 25 30
Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu
35 40 45
Asn Asn Tyr Lys Thr Thr Pro Pro Val Leu Asp Ser Asp Gly Ser Phe
50 55 60
Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln Gln Gly
65 70 75 80

Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr  
85 90 95

Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys  
100 105

<210> 278

<400> 278  
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<210> 279

<400> 279  
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<210> 280

<400> 280  
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<210> 281

<400> 281  
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<210> 282

<400> 282  
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<210> 283

<400> 283  
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<210> 284

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<210> 285

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<210> 294

<400> 294  
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<210> 295

<400> 295  
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<210> 296

<400> 296  
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<210> 297  
<211> 1464  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CART E6 CD8/CD8/CD28/CD3z sequence - NCD81shuMNE6scFvCD8ecd  
fragment CD8 transmembrane CD28 CD3zetaC

<400> 297  
atggccctgc ccgtgaccgc ttgtctgctc cccctggcgc tgctgctgca cgccgccagg 60  
ccagagggtcc agctgggtga gagtggcggt gggctgggta agcctggcgg ctccctgcgg 120  
ctgagctgcg ccgcgagtgg atttactttc agccgatatg ggatgagttg ggtgcggcaa 180  
gctcccgga agaggctgga atgggtctca acaatctccg gggggggcac ttacatctat 240  
taccgact cagtcaagg gagatttacc atttcacgag acaacgctaa gaataccctg 300  
tatttgcaga tgaattctct gagagcagag gacacagctg ttactattg taccgcgac 360  
aactatggca ggaactacga ctacggtatg gactattggg gacaaggac attggttaca 420  
gtgagcagtg gcggcggggg cagcggagga ggaggcagcg gtgggggggg cagcgagata 480  
gtgctcacgc agtcacccgc gactctcagt ctctcacctg gggaacgagc taccctgacg 540  
tgctctgcta ctcctcagt gtcatatatt cactgggtatc agcaacggcc cgggcagtcc 600  
cctagattgc tcatttatag tacctctaatt ctggcctcag gtatccctgc acgattttct 660  
gcatctggtt caggttctga ttacaccctc actatctcta gcctggagcc tgaagacttt 720

gccgtttatt actgccagca gaggtctagc tccccattca cctttgggag tgggaccaag 780  
gttgaaatta aaacgacaac cccggccccc agaccaccaa cgccagcccc caccatcgcc 840  
agccaacccc tgtctctgag accagaagcc tgtaggcctg ccgccggtgg agctgtgcac 900  
acaagaggac tggatttcgc ctgtgatatc tacatttggg ccccgctcgc aggcacatgt 960  
ggagtgtctc tcctctccct ggtgattacc ctgtactgca gaagcaagcg gtctcggctc 1020  
ctgcattctg attaccatgaa catgacccca agaagaccag gccccaccag gaaacattac 1080  
cagccctacg ctccgccacg cgacttcgct gcctaccggt cccgcgttaa gttctcccga 1140  
tcagccgacg cgcctgctta caagcagggc cagaaccaac tgtacaacga gctgaatctc 1200  
ggtagacggg aagagtacga cgtgttggac aaacggagag gccgcgaccc agaaatgggc 1260  
ggcaagcctc gcaggaaaaa cccccaggag ggactgtaca atgagttgca gaaagataag 1320  
atggcagaag cttatagcga gatcggaatg aagggggaaa ggagacgagg gaaaggacac 1380  
gacggccttt atcagggcct gtccacagca aaaaaagata cgtatgacgc cctccatatg 1440  
caggcacttc caccacggtg ataa 1464

<210> 298  
<211> 486  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> CART E6 CD8/CD8/CD28/CD3z sequence - NCD8lshuMNE6scFvCD8ecd  
fragment CD8 transmembrane CD28 CD3zetaC  
  
<400> 298

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15  
  
His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30  
  
Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45  
  
Thr Phe Ser Arg Tyr Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60  
  
Arg Leu Glu Trp Val Ser Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr  
65 70 75 80  
  
Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95  
  
Lys Asn Thr Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110  
  
Ala Val Tyr Tyr Cys Thr Arg Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr  
115 120 125

Gly Met Asp Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly  
 130 135 140

Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Ile  
 145 150 155 160

Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly Glu Arg  
 165 170 175

Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp  
 180 185 190

Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr Ser Thr  
 195 200 205

Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser  
 210 215 220

Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe  
 225 230 235 240

Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly  
 245 250 255

Ser Gly Thr Lys Val Glu Ile Lys Thr Thr Thr Pro Ala Pro Arg Pro  
 260 265 270

Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser Leu Arg Pro  
 275 280 285

Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr Arg Gly Leu  
 290 295 300

Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr Cys  
 305 310 315 320

Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys Arg Ser Lys  
 325 330 335

Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met Thr Pro Arg Arg  
 340 345 350

Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala Pro Pro Arg Asp  
 355 360 365

Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser Arg Ser Ala Asp Ala  
 370 375 380

Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu Leu Asn Leu  
 385 390 395 400

Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg Gly Arg Asp  
405 410 415

Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln Glu Gly Leu  
420 425 430

Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Tyr Ser Glu Ile  
435 440 445

Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp Gly Leu Tyr  
450 455 460

Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr Asp Ala Leu His Met  
465 470 475 480

Gln Ala Leu Pro Pro Arg  
485

<210> 299

<400> 299  
000

<210> 300  
<211> 1467  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CART E6 CD8/CD8/41BB/CD3z sequence - NCD8lshuMNE6scFvCD8ecd  
fragment CD8 transmembrane 41BB CD3zetaC

<400> 300  
atggccctgc ccgtgaccgc tttgtgctc cccctggcgc tgctgctgca cgccgccagg 60  
ccagaggtcc agctggttga gagtggcggg gggctggtta agcctggcgg ctccctgcgg 120  
ctgagctgcg ccgcgagtgg atttactttc agccgatatg ggatgagttg ggtgcggcaa 180  
gctcccgga agaggctgga atgggtctca acaatctccg gggggggcac ttacatctat 240  
taccgcgact cagtcaaggg gagatttacc atttcacgag acaacgctaa gaataccctg 300  
tatttgaga tgaattctct gagagcagag gacacagctg ttactattg taccgcgac 360  
aactatggca ggaactacga ctacggatat gactattggg gacaaggac attggttaca 420  
gtgagcagtg gcggcggggg cagcggagga ggaggcagcg gtgggggggg cagcgagata 480  
gtgctcacgc agtcacccgc gactctcagt ctctcacctg gggaacgagc taccctgacg 540  
tgctctgcta ctcctcagt gtcatatatt cactggatat agcaacggcc cgggcagtcc 600  
cctagattgc tcatttatag tacctctaatt ctggcctcag gtatccctgc acgattttct 660  
ggatctgggt caggttctga ttacaccctc actatctcta gcctggagcc tgaagacttt 720  
gccgtttatt actgccagca gaggtctagc tccccattca cttttgggag tgggaccaag 780  
gttgaaatta aaacgacaac cccggccccc agaccaccaa cgccagcccc caccatcgcc 840  
agccaacccc tgtctctgag accagaagcc tgtaggcctg ccgccgtgg agctgtgcac 900

```

acaagaggac tggatttcgc ctgtgatatc tacatttggg ccccgctcgc aggcacatgt      960
ggagtgtctcc tcctctccct ggtgattacc ctgtactgca aaaggggccg caaaaaactc      1020
ctttacattt ttaagcagcc ttttatgagg ccagtacaga cgactcaaga ggaagacggg      1080
tgctcatgcc gctttcctga ggaggaggaa ggagggtgcg aactgcgcgt taagttctcc      1140
cgatcagccg acgcgcctgc ttacaagcag ggccagaacc aactgtacaa cgagctgaat      1200
ctcggtagac gggaagagta cgacgtgttg gacaaacgga gaggccgcga cccagaaatg      1260
ggcggcaagc ctcgcaggaa aaacccccag gagggactgt acaatgagtt gcagaaagat      1320
aagatggcag aagcttatag cgagatcgga atgaaggggg aaaggagacg agggaaagga      1380
cacgacggcc tttatcaggg cctgtccaca gcaacaaaag atacgtatga cgccctccat      1440
atgcaggcac ttccaccacg gtgataa      1467

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<210> 301

<211> 487

<212> PRT

<213> Artificial Sequence

<220>

<223> CART E6 CD8/CD8/41BB/CD3z sequence - NCD81shuMNE6scFvCD8ecd  
fragment CD8 transmembrane 41BB CD3zetaC

<400> 301

```

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu
1           5           10          15

```

```

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu
          20          25          30

```

```

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe
          35          40          45

```

```

Thr Phe Ser Arg Tyr Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys
          50          55          60

```

```

Arg Leu Glu Trp Val Ser Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr
          65          70          75          80

```

```

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala
          85          90          95

```

```

Lys Asn Thr Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr
          100          105          110

```

```

Ala Val Tyr Tyr Cys Thr Arg Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr
          115          120          125

```

```

Gly Met Asp Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly
          130          135          140

```

Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Ile  
145 150 155 160

Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly Glu Arg  
165 170 175

Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp  
180 185 190

Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr Ser Thr  
195 200 205

Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser  
210 215 220

Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe  
225 230 235 240

Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly  
245 250 255

Ser Gly Thr Lys Val Glu Ile Lys Thr Thr Thr Pro Ala Pro Arg Pro  
260 265 270

Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser Leu Arg Pro  
275 280 285

Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr Arg Gly Leu  
290 295 300

Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr Cys  
305 310 315 320

Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys Lys Arg Gly  
325 330 335

Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe Met Arg Pro Val  
340 345 350

Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe Pro Glu Glu  
355 360 365

Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser Arg Ser Ala Asp  
370 375 380

Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu Leu Asn  
385 390 395 400

Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg Gly Arg  
405 410 415

Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln Glu Gly



420

425

430

Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Tyr Ser Glu  
435 440 445

Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp Gly Leu  
450 455 460

Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr Asp Ala Leu His  
465 470 475 480

Met Gln Ala Leu Pro Pro Arg  
485

<210> 302

<400> 302  
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<210> 303

<211> 1590

<212> DNA

<213> Artificial Sequence

<220>

<223> CART E6 CD8/CD8/CD28/41BB/CD3z sequence - NCD81shuMNE6scFvCD8ecd  
fragment CD8 transmembrane CD28 41BB CD3zetaC

<400> 303

atggccctgc ccgtgaccgc tttgctgctc cccctggcgc tgctgctgca cgccgccagg	60
ccagaggctcc agctgggttga gaggggcggg gggctgggtta agcctggcgg ctccctgcgg	120
ctgagctgcg ccgagagtgg atttactttc agccgatatg ggatgagttg ggtgcggcaa	180
gctcccgga agaggctgga atgggtctca acaatctccg gggggggcac ttacatctat	240
taccccgact cagtcaaggg gagatttacc atttcacgag acaacgctaa gaataccctg	300
tatttgcaga tgaattctct gagagcagag gacacagctg ttactattg taccgagac	360
aactatggca ggaactacga ctacggatatg gactattggg gacaaggac attggttaca	420
gtgagcagtg gcggcggggg cagcggagga ggaggcagcg gtgggggggg cagcgagata	480
gtgctcacgc agtcaccgc gactctcagt ctctcacctg gggaacgagc taccctgacg	540
tgctctgcta cctcctcagt gtcatatatt cactggatat agcaacggcc cgggcagtcc	600
cctagattgc tcatttatag tacctctaatt ctggcctcag gtatccctgc acgattttct	660
ggatctgggt caggttctga ttacaccctc actatctcta gcctggagcc tgaagacttt	720
gccgtttatt actgccagca gaggtctagc tccccattca ctttgggag tgggaccaag	780
gttgaaatta aaacgacaac cccggcccc agaccaccaa cgccagcccc caccatcgcc	840
agccaacccc tgtctctgag accagaagcc tgtaggcctg ccgccggtgg agctgtgcac	900
acaagaggac tggatttcgc ctgtgatatc tacatttggg ccccgctcgc aggacatgt	960
ggagtgtccc tcctctccct ggtgattacc ctgtactgca gaagcaagcg gtctcggctc	1020
ctgcattctg attacatgaa catgacccca agaagaccag gccccaccag gaaacattac	1080

cagccctacg ctccgccacg cgacttcgct gcctaccggt ccaaaagggg ccgcaaaaaa	1140
ctcctttaca tttttaagca gccttttatg aggccagtac agacgactca agaggaagac	1200
gggtgctcat gccgctttcc tgaggaggag gaaggagggt gcgaactgcg cgттаagttc	1260
tccccgatcag ccgacgcgcc tgcttacaag cagggccaga accaactgta caacgagctg	1320
aatctcggta gacgggaaga gtacgacgtg ttggacaaac ggagaggccg cgacccagaa	1380
atgggcggca agcctcgcag gaaaaacccc caggagggac tgtacaatga gttgcagaaa	1440
gataagatgg cagaagctta tagcgagatc ggaatgaagg gggaaaggag acgagggaaa	1500
ggacacgacg gcctttatca gggcctgtcc acagcaacaa aagatacgta tgacgccctc	1560
catatgcagg cacttccacc acggtgataa	1590

<210> 304

<211> 528

<212> PRT

<213> Artificial Sequence

<220>

<223> CART E6 CD8/CD8/CD28/41BB/CD3z sequence - NCD8lshuMNE6scFvCD8ecd  
fragment CD8 transmembrane CD28 41BB CD3zetaC

<400> 304

Met	Ala	Leu	Pro	Val	Thr	Ala	Leu	Leu	Leu	Pro	Leu	Ala	Leu	Leu	Leu
1				5				10					15		

His	Ala	Ala	Arg	Pro	Glu	Val	Gln	Leu	Val	Glu	Ser	Gly	Gly	Gly	Leu
			20					25					30		

Val	Lys	Pro	Gly	Gly	Ser	Leu	Arg	Leu	Ser	Cys	Ala	Ala	Ser	Gly	Phe
		35					40					45			

Thr	Phe	Ser	Arg	Tyr	Gly	Met	Ser	Trp	Val	Arg	Gln	Ala	Pro	Gly	Lys
50						55					60				

Arg	Leu	Glu	Trp	Val	Ser	Thr	Ile	Ser	Gly	Gly	Gly	Thr	Tyr	Ile	Tyr
65					70					75				80	

Tyr	Pro	Asp	Ser	Val	Lys	Gly	Arg	Phe	Thr	Ile	Ser	Arg	Asp	Asn	Ala
				85					90					95	

Lys	Asn	Thr	Leu	Tyr	Leu	Gln	Met	Asn	Ser	Leu	Arg	Ala	Glu	Asp	Thr
		100						105					110		

Ala	Val	Tyr	Tyr	Cys	Thr	Arg	Asp	Asn	Tyr	Gly	Arg	Asn	Tyr	Asp	Tyr
		115					120					125			

Gly	Met	Asp	Tyr	Trp	Gly	Gln	Gly	Thr	Leu	Val	Thr	Val	Ser	Ser	Gly
130						135					140				

Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser	Glu	Ile
145					150					155				160	

Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly Glu Arg  
165 170 175

Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp  
180 185 190

Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr Ser Thr  
195 200 205

Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser  
210 215 220

Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe  
225 230 235 240

Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly  
245 250 255

Ser Gly Thr Lys Val Glu Ile Lys Thr Thr Thr Pro Ala Pro Arg Pro  
260 265 270

Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser Leu Arg Pro  
275 280 285

Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr Arg Gly Leu  
290 295 300

Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr Cys  
305 310 315 320

Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys Arg Ser Lys  
325 330 335

Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met Thr Pro Arg Arg  
340 345 350

Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala Pro Pro Arg Asp  
355 360 365

Phe Ala Ala Tyr Arg Ser Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile  
370 375 380

Phe Lys Gln Pro Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp  
385 390 395 400

Gly Cys Ser Cys Arg Phe Pro Glu Glu Glu Gly Gly Cys Glu Leu  
405 410 415

Arg Val Lys Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly  
420 425 430

Gln Asn Gln Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr  
435 440 445

Asp Val Leu Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys  
450 455 460

Pro Arg Arg Lys Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys  
465 470 475 480

Asp Lys Met Ala Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg  
485 490 495

Arg Arg Gly Lys Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala  
500 505 510

Thr Lys Asp Thr Tyr Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
515 520 525

<210> 305

<400> 305  
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<210> 306  
<211> 1608  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CART C2 CD8/CD8/CD28/41BB/CD3z sequence - NCD8lshuMNC2scFvCD8ecd  
fragment CD8 transmembrane CD28 41BB CD3zetaC

<400> 306  
atggccttgc cagtgcgggc cctgctgctg ccattggctc ttctgttgca cgctgccagg 60  
cctgaagtgc agctcgtaga gagtggcggg ggactggtga agcccgttg aagcctcaga 120  
ctcagttgcg ccgcctcagg tttcactttt tcaggttacg ccatgtcctg ggtaagacag 180  
gcaccgggga aaggactcga gtgggtgtct actatcagct caggaggcac ttatatatat 240  
tatcctgact ctgtaaaagg ccgatttacg atttctcgcg acaatgcaa gaactccctc 300  
tacctccaaa tgaacagtct tagggcagaa gacactgctg tatactattg tgcacgcctc 360  
ggcggcgaca actactacga gtactttgac gtgtggggga aagggactac cgtgacagtt 420  
tcaagcggag gaggtggctc aggtggaggc gggtcagggg ggggaggaag tgatattgtg 480  
ctcacacaat cccagcctc cctggctgtg tctcccgcc aacgcgtac aattacatgt 540  
cgggcctcca aaagcgtgag caccagcggc tacagctaca tgcactggta tcaacagaaa 600  
ccaggacaac ccccaaact gttgatttat ctgcgttcaa acttgagtc cggcgtgcct 660  
gcgcgctttt caggagtggt gagcggcaca gattttacgc tgactatcaa cccgtagaa 720  
gcaaacgata cagcgaatta ttattgtcaa cattcccggg aactcccctt tacgttcggc 780  
gggggcacaa aggtcgaaat taagagaacc acgacaaccc cggccccag accaccaacg 840  
ccagcccca ccatcgccag ccaaccctg tctctgagac cagaagcctg taggcctgcc 900

gccggtggag ctgtgcacac aagaggactg gatttcgcct gtgatatcta catttgggcc 960  
 ccgctcgcag gcacatgtgg agtgctcctc ctctccctgg tgattaccct gtactgcaga 1020  
 agcaagcggc ctcggctcct gcattctgat tacatgaaca tgacccaag aagaccaggc 1080  
 cccaccagga aacattacca gccctacgct ccgccacgcg acttcgctgc ctaccggtcc 1140  
 aaaaggggcc gcaaaaaact cttttacatt tttaagcagc cttttatgag gccagtacag 1200  
 acgactcaag aggaagacgg gtgctcatgc cgctttcctg aggaggagga aggagggtgc 1260  
 gaactgcgcg ttaagttctc ccgatcagcc gacgcgcctg cttacaagca gggccagaac 1320  
 caactgtaca acgagctgaa tctcggtaga cgggaagagt acgacgtgtt ggacaaacgg 1380  
 agaggccgcg acccagaaat gggcggcaag cctcgcagga aaaaccccca ggagggactg 1440  
 tacaatgagt tgcagaaaga taagatggca gaagcttata gcgagatcgg aatgaagggg 1500  
 gaaaggagac gagggaaagg acacgacggc ctttatcagg gcctgtccac agcaacaaaa 1560  
 gatacgtatg acgccctcca tatgcaggca cttccaccac ggtgataa 1608

<210> 307

<211> 513

<212> PRT

<213> Artificial Sequence

<220>

<223> CART C2 CD8/CD8/CD28/41BB/CD3z sequence - NCD81shuMNC2scFvCD8ecd  
fragment CD8 transmembrane CD28 41BB CD3zetaC

<400> 307

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
 1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
 20 25 30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
 35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
 50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
 65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
 85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val Trp Gly  
 100 105 110

Lys Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly  
 115 120 125

Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Leu Thr Gln Ser Pro  
 130 135 140

Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala Thr Ile Thr Cys Arg  
 145 150 155 160

Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser Tyr Met His Trp Tyr  
 165 170 175

Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Leu Ala Ser  
 180 185 190

Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly  
 195 200 205

Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu Ala Asn Asp Thr Ala  
 210 215 220

Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro Phe Thr Phe Gly Gly  
 225 230 235 240

Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr Thr Pro Ala Pro Arg  
 245 250 255

Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser Leu Arg  
 260 265 270

Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr Arg Gly  
 275 280 285

Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr  
 290 295 300

Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys Arg Ser  
 305 310 315 320

Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met Thr Pro Arg  
 325 330 335

Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala Pro Pro Arg  
 340 345 350

Asp Phe Ala Ala Tyr Arg Ser Lys Arg Gly Arg Lys Lys Leu Leu Tyr  
 355 360 365

Ile Phe Lys Gln Pro Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu  
 370 375 380

Asp Gly Cys Ser Cys Arg Phe Pro Glu Glu Glu Glu Gly Gly Cys Glu  
 385 390 395 400

Leu Arg Val Lys Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln

405

410

415

Gly Gln Asn Gln Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu  
420 425 430

Tyr Asp Val Leu Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly  
435 440 445

Lys Pro Arg Arg Lys Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln  
450 455 460

Lys Asp Lys Met Ala Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu  
465 470 475 480

Arg Arg Arg Gly Lys Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr  
485 490 495

Ala Thr Lys Asp Thr Tyr Asp Ala Leu His Met Gln Ala Leu Pro Pro  
500 505 510

Arg

&lt;210&gt; 308

<400> 308  
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&lt;210&gt; 309

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<210> 340

<400> 340  
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<210> 341

<211> 729

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized E6 scFV sequence in CAR

<400> 341

gaggtccagc tggttgagag tggcgggtggg ctggttaagc ctggcggctc cctgcggctg



Pro Ala Thr Leu Ser Leu Ser Pro Gly Glu Arg Ala Thr Leu Thr Cys  
145 150 155 160

Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp Tyr Gln Gln Arg Pro  
165 170 175

Gly Gln Ser Pro Arg Leu Leu Ile Tyr Ser Thr Ser Asn Leu Ala Ser  
180 185 190

Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Ser Asp Tyr Thr  
195 200 205

Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys  
210 215 220

Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly Ser Gly Thr Lys Val  
225 230 235 240

Glu Ile Lys

<210> 343  
<211> 63  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 leader sequence

<400> 343  
atggccctgc ccgtgaccgc tttgctgctc cccctggcgc tgctgctgca cgccgccagg 60  
cca 63

<210> 344  
<211> 21  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CD8 leader sequence

<400> 344

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro  
20

<210> 345  
<211> 135  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 hinge domain sequence

<400> 345  
acgacaaccc cggccccag accaccaacg ccagcccca ccatgccag ccaaccctg 60  
tctctgagac cagaagcctg taggcctgcc gccggtggag ctgtgcacac aagaggactg 120  
gatttcgcct gtgat 135

<210> 346  
<211> 45  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CD8 hinge domain sequence

<400> 346  
Thr Thr Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala  
1 5 10 15

Ser Gln Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly  
20 25 30

Gly Ala Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp  
35 40 45

<210> 347  
<211> 66  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD4 hinge domain sequence

<400> 347  
tcgggacagg tcctgctgga atccaacatc aaggttctgc ccacatggtc caccgcgtg 60  
cagcca 66

<210> 348  
<211> 22  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CD4 hinge domain sequence

<400> 348  
Ser Gly Gln Val Leu Leu Glu Ser Asn Ile Lys Val Leu Pro Thr Trp  
1 5 10 15

Ser Thr Pro Val Gln Pro  
20

<210> 349  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>

<223> CD28 hinge domain sequence

<400> 349

aaacaccttt gtccaagtcc cctatttccc ggaccttcta agccc

45

<210> 350

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> CD28 hinge domain sequence

<400> 350

Lys His Leu Cys Pro Ser Pro Leu Phe Pro Gly Pro Ser Lys Pro  
1 5 10 15

<210> 351

<400> 351  
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<210> 360

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<210> 361  
 <211> 63  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> CD3 zeta transmembrane domain sequence  
  
 <400> 361  
 ctctgctacc tgctggatgg aatcctcttc atctatggtg tcattctcac tgccttggtc 60  
  
 ctg 63

<210> 362  
 <211> 21  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> CD3 zeta transmembrane domain sequence  
  
 <400> 362  
  
 Leu Cys Tyr Leu Leu Asp Gly Ile Leu Phe Ile Tyr Gly Val Ile Leu  
 1 5 10 15  
  
 Thr Ala Leu Phe Leu  
 20

<210> 363  
 <211> 72  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> CD8 transmembrane domain sequence  
  
 <400> 363  
 atctacattt gggccccgct cgcaggcaca tgtggagtgc tcctcctctc cctgggtgatt 60  
  
 accctgtact gc 72

<210> 364  
 <211> 24  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> CD8 transmembrane domain sequence  
  
 <400> 364  
  
 Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu  
 1 5 10 15  
  
 Ser Leu Val Ile Thr Leu Tyr Cys  
 20

<210> 365  
 <211> 66  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD4 transmembrane domain sequence

<400> 365  
 atggccctga ttgtgctggg gggcgtcgcc ggcctcctgc ttttcattgg gctaggcatc 60  
 ttcttc 66

<210> 366  
 <211> 22  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> CD4 transmembrane domain sequence

<400> 366

Met Ala Leu Ile Val Leu Gly Gly Val Ala Gly Leu Leu Leu Phe Ile  
 1 5 10 15

Gly Leu Gly Ile Phe Phe  
 20

<210> 367  
 <211> 81  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD28 transmembrane domain sequence

<400> 367  
 ttttgggtgc tgggtgggtg tgggtggagtc ctggcttgct atagcttgct agtaacagtg 60  
 gcctttatta ttttctgggt g 81

<210> 368  
 <211> 27  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> CD28 transmembrane domain sequence

<400> 368

Phe Trp Val Leu Val Val Val Gly Gly Val Leu Ala Cys Tyr Ser Leu  
 1 5 10 15

Leu Val Thr Val Ala Phe Ile Ile Phe Trp Val  
 20 25

<210> 369  
 <211> 81  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> 41BB transmembrane domain sequence

<400> 369

atcatctcct tctttcttgc gctgacgtcg actgcgttgc tcttcctgct gttcttcctc 60  
acgctccggt tctctgttgt t 81



gagttgcaga aagataagat ggcagaagct tatagcgaga tcggaatgaa gggggaaagg	240
agacgaggga aaggacacga cggcctttat cagggcctgt ccacagcaac aaaagatacg	300
tatgacgccc tccatatgca ggcacttcca ccacgg	336

<210> 374  
 <211> 112  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> CD3 zeta domain sequence

<400> 374

Arg Val Lys Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly	
1                  5                  10                  15	

Gln Asn Gln Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr	
20                  25                  30	

Asp Val Leu Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys	
35                  40                  45	

Pro Arg Arg Lys Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys	
50                  55                  60	

Asp Lys Met Ala Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg	
65                  70                  75                  80	

Arg Arg Gly Lys Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala	
85                  90                  95	

Thr Lys Asp Thr Tyr Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg	
100                  105                  110	

<210> 375  
 <211> 336  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD3 zeta domain variant sequence

<400> 375

agagtgaagt tcagcaggag cgcagacgcc cccgcgtacc agcagggccca gaaccagctc	60
tataacgagc tcaatctagg acgaagagag gagtacgatg ttttggacaa gagacgtggc	120
cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgtacaat	180
gaactgcaga aagataagat ggcggaggcc tacagtgaga ttgggatgaa aggcgagcgc	240
cggagggggca aggggcacga tggcctttac caggggtctca gtacagccac caaggacacc	300
tacgacgccc ttcacatgca ggccctgccc cctcgc	336

<210> 376

<211> 112  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CD3 zeta domain variant sequence

<400> 376

Arg Val Lys Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Gln Gln Gly  
1 5 10 15

Gln Asn Gln Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr  
20 25 30

Asp Val Leu Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys  
35 40 45

Pro Arg Arg Lys Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys  
50 55 60

Asp Lys Met Ala Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg  
65 70 75 80

Arg Arg Gly Lys Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala  
85 90 95

Thr Lys Asp Thr Tyr Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
100 105 110

<210> 377  
<211> 123  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD28 domain sequence

<400> 377

agaagcaagc ggtctcggct cctgcattct gattacatga acatgacccc aagaagacca 60

ggccccacca ggaaacatta ccagccctac gctccgccac gcgacttcgc tgcctaccgg 120

tcc 123

<210> 378  
<211> 41  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CD28 domain sequence

<400> 378

Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met Thr  
1 5 10 15

Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala Pro  
20 25 30

Pro Arg Asp Phe Ala Ala Tyr Arg Ser  
35 40

<210> 379  
<211> 126  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> 41BB domain sequence

<400> 379  
aaaaggggcc gcaaaaaact cctttacatt tttaagcagc cttttatgag gccagtacag 60  
acgactcaag aggaagacgg gtgctcatgc cgctttcctg aggaggagga aggagggtgc 120  
gaactg 126

<210> 380  
<211> 42  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> 41BB domain sequence

<400> 380

Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe Met  
1 5 10 15

Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe  
20 25 30

Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu  
35 40

<210> 381

<400> 381  
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<210> 382

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<210> 390

<400> 390  
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<210> 391

<211> 729

<212> DNA

<213> Artificial Sequence

<220>

<223> HumanizeE6 scFV (VHVL) sequence

<400> 391

gaggtgcagc tggaggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc	60
tcctgtgcag cctctggatt caccttcagt aggtatggca tgagctgggt ccgccaggct	120
ccagggaaga ggctggagtg ggtctcaacc attagtggcg gaggcaccta catatactac	180
ccagactcag tgaagggccg attcaccatc tccagagaca acgccaagaa caccctgtat	240
ctgcaaatga acagcctgag agccgaggac acggctgtgt attactgtac cagagataac	300
tatggccgca actatgatta tggcatggat tattggggcc agggcaccct ggtgaccgtg	360
agcagcggcg gtggcggatc cggcgggtggc ggatccggcg gtggcggatc cgaaattgtg	420
ttgacacagt ctccagccac cctgtctttg tctccagggg aaagagccac cctcacctgc	480
agcgccacca gcagtgttag ctacatccac tggtagcaac agaggcctgg ccagagcccc	540
aggctcctca tctatagcac ctccaacctg gccagcggca tcccagccag gttcagtggc	600
agtgggtctg ggagcgacta cactctcacc atcagcagcc tagagcctga agattttgca	660
gtttattact gtcagcagcg tagcagctcc cctttcacct ttggcagcgg caccaaagtg	720
gaaattaaa	729

<210> 392

<211> 243

<212> PRT

<213> Artificial Sequence

<220>

<223> HumanizeE6 scFV (VHVL) sequence

<400> 392

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Arg Tyr  
20 25 30

Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys Arg Leu Glu Trp Val  
35 40 45

Ser Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr  
65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Thr Arg Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr Gly Met Asp Tyr Trp  
100 105 110

Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly  
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Ile Val Leu Thr Gln Ser  
130 135 140

Pro Ala Thr Leu Ser Leu Ser Pro Gly Glu Arg Ala Thr Leu Thr Cys  
145 150 155 160

Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp Tyr Gln Gln Arg Pro  
165 170 175

Gly Gln Ser Pro Arg Leu Leu Ile Tyr Ser Thr Ser Asn Leu Ala Ser  
180 185 190

Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Ser Asp Tyr Thr  
195 200 205

Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys  
210 215 220

Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly Ser Gly Thr Lys Val  
225 230 235 240

Glu Ile Lys

<210> 393

<211> 729

<212> DNA

<213> Artificial Sequence

<220>

<223> HumanizeE6 scFV (VLVH) sequence

<400> 393

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gaaattgtgt tgacacagtc tccagccacc ctgtctttgt ctccagggga aagagccacc      60
ctcacctgca gcgccaccag cagtgttagc tacatccact ggtaccaaca gaggcctggc      120
cagagcccca ggctcctcat ctatagcacc tccaacctgg ccagcggcat cccagccagg      180
ttcagtggca gtgggtctgg gagcgactac actctcacca tcagcagcct agagcctgaa      240
gattttgcag tttattactg tcagcagcgt agcagctccc ctttcacctt tggcagcggc      300
accaaagtgg aaattaaagg cgggtggcgga tccggcggtg gcggatccgg cggtggcgga      360
tccgaggtgc agctgggtgga gtctggggga ggcctggtca agcctggggg gtccctgaga      420
ctctcctgtg cagcctctgg attcaccttc agtaggtatg gcatgagctg ggtccgccag      480
gctccaggga agaggctgga gtgggtctca accattagtg gcggaggcac ctacatatac      540
taccagact cagtgaaggg ccgattcacc atctccagag acaacgcaa gaacaccctg      600
tatctgcaaa tgaacagcct gagagccgag gacacggctg tgtattactg taccagagat      660
aactatggcc gcaactatga ttatggcatg gattattggg gccagggcac cctggtgacc      720
gtgagcagc                                     729
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<210> 394

<211> 243

<212> PRT

<213> Artificial Sequence

<220>

<223> HumanizeE6 scFV (VLVH) sequence

<400> 394

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Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly
1              5              10             15
```

```
Glu Arg Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile
20              25             30
```

```
His Trp Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr
35              40             45
```

```
Ser Thr Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser
50              55             60
```

```
Gly Ser Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu
65              70             75             80
```

```
Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr
85              90             95
```

```
Phe Gly Ser Gly Thr Lys Val Glu Ile Lys Gly Gly Gly Gly Ser Gly
100             105            110
```

Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Val Gln Leu Val Glu Ser  
115 120 125

Gly Gly Gly Leu Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala  
130 135 140

Ala Ser Gly Phe Thr Phe Ser Arg Tyr Gly Met Ser Trp Val Arg Gln  
145 150 155 160

Ala Pro Gly Lys Arg Leu Glu Trp Val Ser Thr Ile Ser Gly Gly Gly  
165 170 175

Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser  
180 185 190

Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln Met Asn Ser Leu Arg  
195 200 205

Ala Glu Asp Thr Ala Val Tyr Tyr Cys Thr Arg Asp Asn Tyr Gly Arg  
210 215 220

Asn Tyr Asp Tyr Gly Met Asp Tyr Trp Gly Gln Gly Thr Leu Val Thr  
225 230 235 240

Val Ser Ser

<210> 395  
<211> 747  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> HumanizeC2 scFV (VHVL) sequence

<400> 395  
gaggtgcagc tggaggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60  
tcctgtgcag cctctggatt caccttcagt ggctatgcc a tgagctgggt ccgccaggct 120  
ccagggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac 180  
cccgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240  
ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagacttggg 300  
ggggataatt actacgaata cttcgatgtc tggggcaaag ggaccacggt caccgtctcc 360  
tccggcgggtg gcggatccgg cggtaggcga tccggcgggt gcggatccga cattgtgctg 420  
accagtctc cagcctcctt ggccgtgtct ccaggacaga gggccaccat cacctgcaga 480  
gccagtaaga gtgtcagtac cagcggatac tcctacatgc actggtatca gcagaaacca 540  
ggacaacctc ctaaactcct gatttacctg gcatccaatc tggagagcgg ggtcccagcc 600  
aggttcagcg gcagtgggtc tgggaccgat ttcaccctca caattaatcc tgttgaagct 660

aatgatactg caaattatta ctgtcagcac agtagggagc tgcctttcac attcggcgga 720  
 gggaccaagg tggagatcaa acgaact 747

<210> 396  
 <211> 249  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> HumanizeC2 scFV (VHVL) sequence

<400> 396

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
 1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
 20 25 30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
 35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
 50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
 65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
 85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val Trp Gly  
 100 105 110

Lys Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly  
 115 120 125

Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Leu Thr Gln Ser Pro  
 130 135 140

Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala Thr Ile Thr Cys Arg  
 145 150 155 160

Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser Tyr Met His Trp Tyr  
 165 170 175

Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Leu Ala Ser  
 180 185 190

Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly  
 195 200 205

Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu Ala Asn Asp Thr Ala  
 210 215 220



Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro Phe Thr Phe Gly Gly  
225 230 235 240

Gly Thr Lys Val Glu Ile Lys Arg Thr  
245

<210> 397  
<211> 747  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> HumanizeE6 scFV (VLVH) sequence

<400> 397  
gacattgtgc tgaccagtc tccagcctcc ttggccgtgt ctccaggaca gagggccacc 60  
atcacctgca gagccagtaa gagtgtcagt accagcggat actcctacat gcaactgttat 120  
cagcagaaac caggacaacc tcctaaactc ctgatttacc tggcatcaa tctggagagc 180  
gggggtcccag ccaggttcag cggcagtggg tctgggaccg atttcaccct cacaattaat 240  
cctgtggaag ctaatgatac tgcaaattat tactgtcagc acagtaggga gctgcctttc 300  
acattcggcg gagggaccaa ggtggagatc aaacgaactg gcggtggcgg atccggcggt 360  
ggcggatccg gcggtggcgg atccgaggtg cagctggtgg agtctggggg aggcctggtc 420  
aagcctgggg ggtccctgag actctcctgt gcagcctctg gattcacctt cagtggctat 480  
gccatgagct gggtcgcga ggctccaggg aaggggctgg agtgggtctc aaccattagt 540  
agtggcgga cctacatata ctaccccgac tcagtgaagg gccgattcac catctccaga 600  
gacaacgcca agaactcact gtatctgcaa atgaacagcc tgagagccga ggacacggcc 660  
gtgtattact gtgcgagact tgggggggat aattactacg aatacttcga tgtctggggc 720  
aaagggacca cggtcaccgt ctcctcc 747

<210> 398  
<211> 249  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> HumanizeE6 scFV (VLVH) sequence

<400> 398

Asp Ile Val Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly  
1 5 10 15

Gln Arg Ala Thr Ile Thr Cys Arg Ala Ser Lys Ser Val Ser Thr Ser  
20 25 30

Gly Tyr Ser Tyr Met His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro  
35 40 45

Lys Leu Leu Ile Tyr Leu Ala Ser Asn Leu Glu Ser Gly Val Pro Ala

50	55	60
Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn 65 70 75 80		
Pro Val Glu Ala Asn Asp Thr Ala Asn Tyr Tyr Cys Gln His Ser Arg 85 90 95		
Glu Leu Pro Phe Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg 100 105 110		
Thr Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser 115 120 125		
Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly 130 135 140		
Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr 145 150 155 160		
Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val 165 170 175		
Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val 180 185 190		
Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr 195 200 205		
Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys 210 215 220		
Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val Trp Gly 225 230 235 240		
Lys Gly Thr Thr Val Thr Val Ser Ser 245		

<210> 399  
 <211> 15  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> G4S1 linker sequence

<400> 399  
 ggcgggtggcg gatcc

<210> 400  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence

<220>

<223> G4S1 linker sequence

<400> 400

Gly Gly Gly Gly Ser  
1 5

<210> 401

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> [G4S1]x3 linker sequence

<400> 401

ggcgggtggcg gatccggcgg tggcggatcc ggcgggtggcg gatcc

45

<210> 402

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> [G4S1]x3 linker sequence

<400> 402

Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser  
1 5 10 15

<210> 403

<211> 24

<212> DNA

<213> Artificial Sequence

<220>

<223> 8 aa GS linker sequence

<400> 403

ggcgggttccg gcggtggatc cgga

24

<210> 404

<211> 8

<212> PRT

<213> Artificial Sequence

<220>

<223> 8 aa GS linker sequence

<400> 404

Gly Gly Ser Gly Gly Gly Ser Gly  
1 5

<210> 405

<211> 39

<212> DNA

<213> Artificial Sequence

<220>

<223> 12 aa GS linker sequence

<400> 405

ggcggttccg gcggtggatc cggcgggtggc ggatccgga

39

<210> 406  
<211> 12  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> 12 aa GS linker sequence

<400> 406

Gly Gly Ser Gly Gly Gly Ser Gly Gly Gly Ser Gly  
1 5 10

<210> 407  
<211> 39  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> 13 aa GS linker sequence

<400> 407

ggcgggtggat ccggcgggtgg cggatccggc ggtggatcc

39

<210> 408  
<211> 13  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> 13 aa GS linker sequence

<400> 408

Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Ser  
1 5 10

<210> 409  
<211> 66  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> 22 aa GS linker sequence

<400> 409

ggcgggtggaa gcggcgggtgg cggatccggc agcggcggaa gcggcgggtgg cggatccggc

60

ggtgga

66

<210> 410  
<211> 22  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> 22 aa GS linker sequence

<400> 410

Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Ser Gly Gly Ser Gly Gly  
1 5 10 15

Gly Gly Ser Gly Gly Gly  
20

<210> 411  
<211> 78  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> 24 aa GS linker sequence

<400> 411  
ggcggttccg gcggtggatc cggcgggtggc ggatccggag gcggttccgg cgggtggatcc 60  
ggcgggtggcg gatccgga 78

<210> 412  
<211> 24  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> 24 aa GS linker sequence

<400> 412

Gly Gly Ser Gly Gly Gly Ser Gly Gly Gly Ser Gly Gly Gly Ser Gly  
1 5 10 15

Gly Gly Ser Gly Gly Gly Ser Gly  
20

<210> 413  
<211> 357  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse C3 Heavy chain variable region sequence

<400> 413  
caggtccagc tgcagcagtc tgggcctgag ctggtgaggc ctgggggtctc agtgaagatt 60  
tcctgcaagg gttccggcta cagattcact gattatgcta tgaactgggt gaagcagagt 120  
catgcaaaga gtctagagtg gattggagtt attagtactt tctctggtaa taaaaacttc 180  
aaccagaagt ttaagggcaa ggccacaatg actgtagaca aatcctccag cacagcctat 240  
atggaacttg ccagattgac atctgaggat tctgccatgt attactgtgc aagatcggat 300  
tactacggcc catactttga ctactggggc caaggcacca ctctcacagt ctcctca 357

<210> 414  
<211> 119  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse C3 Heavy chain variable region sequence

<400> 414

Gln Val Gln Leu Gln Gln Ser Gly Pro Glu Leu Val Arg Pro Gly Val  
1 5 10 15

Ser Val Lys Ile Ser Cys Lys Gly Ser Gly Tyr Arg Phe Thr Asp Tyr  
20 25 30

Ala Met Asn Trp Val Lys Gln Ser His Ala Lys Ser Leu Glu Trp Ile  
35 40 45

Gly Val Ile Ser Thr Phe Ser Gly Asn Thr Asn Phe Asn Gln Lys Phe  
50 55 60

Lys Gly Lys Ala Thr Met Thr Val Asp Lys Ser Ser Ser Thr Ala Tyr  
65 70 75 80

Met Glu Leu Ala Arg Leu Thr Ser Glu Asp Ser Ala Met Tyr Tyr Cys  
85 90 95

Ala Arg Ser Asp Tyr Tyr Gly Pro Tyr Phe Asp Tyr Trp Gly Gln Gly  
100 105 110

Thr Thr Leu Thr Val Ser Ser  
115

<210> 415

<211> 90

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C3 heavy chain variable framework region 1 (FWR1) sequence

<400> 415

cagggtccagc tgcagcagtc tgggcctgag ctggtgaggc ctgggggtctc agtgaagatt 60

tcctgcaagg gttccggcta cagattcact 90

<210> 416

<211> 30

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C3 heavy chain variable framework region 1 (FWR1) sequence

<400> 416

Gln Val Gln Leu Gln Gln Ser Gly Pro Glu Leu Val Arg Pro Gly Val  
1 5 10 15

Ser Val Lys Ile Ser Cys Lys Gly Ser Gly Tyr Arg Phe Thr  
20 25 30

<210> 417

<211> 15

<212> DNA

<213> Artificial Sequence

<220>  
 <223> Mouse C3 heavy chain variable complementarity determining regions  
 1 (CDR1) sequence  
 <400> 417  
 gattatgcta tgaac 15

<210> 418  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse C3 heavy chain variable complementarity determining regions  
 1 (CDR1) sequence

<400> 418

Asp Tyr Ala Met Asn  
 1 5

<210> 419  
 <211> 42  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse C3 heavy chain variable framework region 2 (FWR2) sequence

<400> 419  
 tgggtgaagc agagtcattgc aaagagtcta gagtggattg ga 42

<210> 420  
 <211> 14  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse C3 heavy chain variable framework region 2 (FWR2) sequence

<400> 420

Trp Val Lys Gln Ser His Ala Lys Ser Leu Glu Trp Ile Gly  
 1 5 10

<210> 421  
 <211> 51  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse C3 heavy chain variable complementarity determining regions  
 2 (CDR2) sequence

<400> 421  
 gttattagta ctttctctgg taatacaaac ttcaaccaga agtttaaggg c 51

<210> 422  
 <211> 17  
 <212> PRT  
 <213> Artificial Sequence

<220>

<223> Mouse C3 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 422

Val	Ile	Ser	Thr	Phe	Ser	Gly	Asn	Thr	Asn	Phe	Asn	Gln	Lys	Phe	Lys
1				5					10					15	

Gly

<210> 423

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C3 heavy chain variable framework region 3 (FWR3) acid  
sequence

<400> 423

aaggccacaa tgactgtaga caaatcctcc agcacagcct atatggaact tgccagattg 60

acatctgagg attctgcat gtattactgt gcaaga 96

<210> 424

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C3 heavy chain variable framework region 3 (FWR3) acid  
sequence

<400> 424

Lys	Ala	Thr	Met	Thr	Val	Asp	Lys	Ser	Ser	Ser	Thr	Ala	Tyr	Met	Glu
1				5					10					15	

Leu	Ala	Arg	Leu	Thr	Ser	Glu	Asp	Ser	Ala	Met	Tyr	Tyr	Cys	Ala	Arg
			20					25					30		

<210> 425

<211> 30

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C3 heavy chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 425

tcggattact acggcccata ctttgactac 30

<210> 426

<211> 10

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C3 heavy chain variable complementarity determining regions  
3 (CDR3) sequence



<400> 426

Ser Asp Tyr Tyr Gly Pro Tyr Phe Asp Tyr  
1 5 10

<210> 427

<400> 427  
000

<210> 428

<400> 428  
000

<210> 429

<400> 429  
000

<210> 430

<400> 430  
000

<210> 431

<400> 431  
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<210> 435

<400> 435  
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<210> 436

<400> 436  
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<210> 437

<400> 437  
000

<210> 438

<400> 438  
000

<210> 439

<211> 357

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C3 heavy chain variable region sequence from IGHV118\*04

<400> 439

```
caggttcagc tgggtcagtc tggagctgag gtgaagaagc ctggggcctc agtgaaggtc      60
tcctgcaagg cttctggtta cacctttacc gactacgcca tgaactgggt gcgacaggcc      120
cctggacaag ggcttgagtg gatgggagtg atcagcacct tcagcggtaa cacaaacttc      180
aaccagaagt tcaagggcag agtcaccatg accacagaca catccacgag cacagcctac      240
atggagctga ggagcctgag atctgacgac acggccgtgt attactgtgc gagaagcgac      300
tactacggcc catacttcga ctactggggc cagggcacca ccctgaccgt gtccagc        357
```

<210> 440

<211> 119

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C3 heavy chain variable region sequence from IGHV118\*04

<400> 440

```
Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ala
1              5              10              15
```

```
Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asp Tyr
20              25              30
```

```
Ala Met Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35              40              45
```

```
Gly Val Ile Ser Thr Phe Ser Gly Asn Thr Asn Phe Asn Gln Lys Phe
50              55              60
```

```
Lys Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
65              70              75              80
```

```
Met Glu Leu Arg Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85              90              95
```

```
Ala Arg Ser Asp Tyr Tyr Gly Pro Tyr Phe Asp Tyr Trp Gly Gln Gly
100              105              110
```

```
Thr Thr Leu Thr Val Ser Ser
115
```

<210> 441

<211> 90

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C3 heavy chain variable framework region 1 (FWR1) acid sequence

<400> 441  
caggttcagc tgggtgcagtc tggagctgag gtgaagaagc ctggggcctc agtgaaggtc 60  
tcctgcaagg cttcttggtta cacctttacc 90

<210> 442  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 heavy chain variable framework region 1 (FWR1) acid  
sequence

<400> 442  
Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ala  
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr  
20 25 30

<210> 443  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 heavy chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 443  
gactacgcc tgaac 15

<210> 444  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 heavy chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 444  
Asp Tyr Ala Met Asn  
1 5

<210> 445  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 heavy chain variable framework region 2 (FWR2) acid  
sequence

<400> 445  
tgggtgagc agggccctgg acaagggtt gagtggatgg ga 42

<210> 446  
<211> 14

<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 heavy chain variable framework region 2 (FWR2) acid  
sequence

<400> 446

Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met Gly  
1 5 10

<210> 447  
<211> 51  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 heavy chain variable complementarity determining  
regions 2 (CDR2) sequence

<400> 447  
gtgatcagca ccttcagcgg taacacaaac ttcaaccaga agttcaaggg c 51

<210> 448  
<211> 17  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 heavy chain variable complementarity determining  
regions 2 (CDR2) sequence

<400> 448

Val Ile Ser Thr Phe Ser Gly Asn Thr Asn Phe Asn Gln Lys Phe Lys  
1 5 10 15

Gly

<210> 449  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 heavy chain variable framework region 3 (FWR3) acid  
sequence

<400> 449  
agagtcacca tgaccacaga cacatccacg agcacagcct acatggagct gaggagcctg 60  
agatctgacg acacggccgt gtattactgt gcgaga 96

<210> 450  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 heavy chain variable framework region 3 (FWR3) acid  
sequence

<400> 450

Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr Met Glu  
1 5 10 15

Leu Arg Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys Ala Arg  
20 25 30

<210> 451

<211> 30

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C3 heavy chain variable complementarity determining  
regions 3 (CDR3) sequence

<400> 451

agcgactact acggcccata cttcgactac 30

<210> 452

<211> 10

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C3 heavy chain variable complementarity determining  
regions 3 (CDR3) sequence

<400> 452

Ser Asp Tyr Tyr Gly Pro Tyr Phe Asp Tyr  
1 5 10

<210> 453

<211> 1353

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C3 IgG1 heavy chain sequence

<400> 453

caggttcagc tgggtgcagtc tggagctgag gtgaagaagc ctggggcctc agtgaaggtc 60

tcctgcaagg cttctggtta cacctttacc gactacgcca tgaactgggt gcgacaggcc 120

cctggacaag ggcttgagt gatgggagt atcagcacct tcagcggtaa cacaaacttc 180

aaccagaagt tcaagggcag agtcaccatg accacagaca catccacgag cacagcctac 240

atggagctga ggagcctgag atctgacgac acggccgtgt attactgtgc gagaagcgac 300

tactacggcc catacttcga ctactggggc cagggcacca ccctgaccgt gtccagcgct 360

agcaccaagg gcccatcggc cttccccctg gcaccctcct ccaagagcac ctctgggggc 420

acagcggccc tgggctgcct ggtcaaggac tacttccccg aaccggtgac ggtgtcgtgg 480

aactcaggcg ccctgaccag cggcgtgcac accttccccg ctgtcctaca gtcctcagga 540

ctctactccc tcagcagcgt ggtgacagt ccctccagca gcttgggcac ccagacctac 600

atctgcaacg tgaatcacia gccagcaac accaaggtgg acaagaaagt tgagcccaaa 660



Gly Cys Leu Val Lys Asp Tyr Phe Pro Glu Pro Val Thr Val Ser Trp  
145 150 155 160

Asn Ser Gly Ala Leu Thr Ser Gly Val His Thr Phe Pro Ala Val Leu  
165 170 175

Gln Ser Ser Gly Leu Tyr Ser Leu Ser Ser Val Val Thr Val Pro Ser  
180 185 190

Ser Ser Leu Gly Thr Gln Thr Tyr Ile Cys Asn Val Asn His Lys Pro  
195 200 205

Ser Asn Thr Lys Val Asp Lys Lys Val Glu Pro Lys Ser Cys Asp Lys  
210 215 220

Thr His Thr Cys Pro Pro Cys Pro Ala Pro Glu Leu Leu Gly Gly Pro  
225 230 235 240

Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser  
245 250 255

Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser His Glu Asp  
260 265 270

Pro Glu Val Lys Phe Asn Trp Tyr Val Asp Gly Val Glu Val His Asn  
275 280 285

Ala Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn Ser Thr Tyr Arg Val  
290 295 300

Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu Asn Gly Lys Glu  
305 310 315 320

Tyr Lys Cys Lys Val Ser Asn Lys Ala Leu Pro Ala Pro Ile Glu Lys  
325 330 335

Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr  
340 345 350

Leu Pro Pro Ser Arg Glu Glu Met Thr Lys Asn Gln Val Ser Leu Thr  
355 360 365

Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu  
370 375 380

Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro Pro Val Leu  
385 390 395 400

Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys  
405 410 415

Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu  
420 425 430

Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly  
435 440 445

Lys

<210> 455  
<211> 1341  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 IgG2 heavy chain sequence

<400> 455  
caggttcagc tgggtgcagtc tggagctgag gtgaagaagc ctggggcctc agtgaaggtc 60  
tcctgcaagg cttctggtta cacctttacc gactacgcca tgaactgggt gcgacaggcc 120  
cctggacaag ggcttgagtg gatgggagtg atcagcacct tcagcggtaa cacaaacttc 180  
aaccagaagt tcaagggcag agtcaccatg accacagaca catccacgag cacagcctac 240  
atggagctga ggagcctgag atctgacgac acggccgtgt attactgtgc gagaagcgac 300  
tactacggcc catacttcga ctactggggc cagggcacca ccctgaccgt gtccagcgcc 360  
tccaccaagg gcccatcggt cttccccctg gcgccctgct ccaggagcac ctccgagagc 420  
acagccgccc tgggctgcct ggtcaaggac tactttcccc aaccgggtgac ggtgtcgtgg 480  
aactcaggcg ctctgaccag ggcgctgcac accttcccag ctgtcctaca gtcctcagga 540  
ctctactccc tcagcagcgt ggtgaccgtg ccctccagca acttcggcac ccagacctac 600  
acctgcaacg tagatcaciaa gcccagcaac accaaggtgg acaagacagt tgagcgcaaa 660  
tgtttgtgtc agtgcccacc gtgcccagca ccacctgtgg caggaccgtc agtcttcctc 720  
ttcccccaa aacccaagga caccctcatg atctcccga cccctgaggt cacgtgcgtg 780  
gtgggtggacg tgagccacga agaccccag gtccagttca actggtacgt ggacggcgtg 840  
gaggtgcata atgccaagac aaagccacgg gaggagcagt tcaacagcac gttccgtgtg 900  
gtcagcgtcc tcaccgttgt gcaccaggac tggctgaacg gcaaggagta caagtgaag 960  
gtctccaaca aaggcctccc agccccatc gagaaaacca tctccaaaac caaagggcag 1020  
ccccgagaac cacaggtgta caccctgccc ccatcccggg aggagatgac caagaaccag 1080  
gtcagcctga cctgcctggt caaaggcttc taccacagcg acatcgccgt ggagtgggag 1140  
agcaatgggc agccggagaa caactacaag accacacctc ccatgctgga ctccgacggc 1200  
tccttcttcc tctacagcaa gtcaccgtg gacaagagca ggtggcagca ggggaacgtc 1260  
ttctcatgct ccgtgatgca tgaggctctg cacaaccact acacgcagaa gaggcctctc 1320  
ctgtctccgg gtaaatagta a 1341



<210> 456  
<211> 445  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 IgG2 heavy chain sequence

<400> 456

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ala  
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asp Tyr  
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met  
35 40 45

Gly Val Ile Ser Thr Phe Ser Gly Asn Thr Asn Phe Asn Gln Lys Phe  
50 55 60

Lys Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr  
65 70 75 80

Met Glu Leu Arg Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Ala Arg Ser Asp Tyr Tyr Gly Pro Tyr Phe Asp Tyr Trp Gly Gln Gly  
100 105 110

Thr Thr Leu Thr Val Ser Ser Ala Ser Thr Lys Gly Pro Ser Val Phe  
115 120 125

Pro Leu Ala Pro Cys Ser Arg Ser Thr Ser Glu Ser Thr Ala Ala Leu  
130 135 140

Gly Cys Leu Val Lys Asp Tyr Phe Pro Glu Pro Val Thr Val Ser Trp  
145 150 155 160

Asn Ser Gly Ala Leu Thr Ser Gly Val His Thr Phe Pro Ala Val Leu  
165 170 175

Gln Ser Ser Gly Leu Tyr Ser Leu Ser Ser Val Val Thr Val Pro Ser  
180 185 190

Ser Asn Phe Gly Thr Gln Thr Tyr Thr Cys Asn Val Asp His Lys Pro  
195 200 205

Ser Asn Thr Lys Val Asp Lys Thr Val Glu Arg Lys Cys Cys Val Glu  
210 215 220

Cys Pro Pro Cys Pro Ala Pro Pro Val Ala Gly Pro Ser Val Phe Leu  
225 230 235 240

Phe Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu  
245 250 255

Val Thr Cys Val Val Val Asp Val Ser His Glu Asp Pro Glu Val Gln  
260 265 270

Phe Asn Trp Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys  
275 280 285

Pro Arg Glu Glu Gln Phe Asn Ser Thr Phe Arg Val Val Ser Val Leu  
290 295 300

Thr Val Val His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys  
305 310 315 320

Val Ser Asn Lys Gly Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys  
325 330 335

Thr Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser  
340 345 350

Arg Glu Glu Met Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys  
355 360 365

Gly Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln  
370 375 380

Pro Glu Asn Asn Tyr Lys Thr Thr Pro Pro Met Leu Asp Ser Asp Gly  
385 390 395 400

Ser Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln  
405 410 415

Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn  
420 425 430

His Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys  
435 440 445

<210> 457

<400> 457  
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<210> 458

<211> 336

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C3 Light Chain variable region sequence

<400> 458

gatgttttga tgacccaaac tccactctcc ctgcctgtca gtcttgaga tcaagcctcc

atctcttgca gatctagtca gaccattgta catagtaatg gaaacaccta tttagaatgg	120
tacctgcaga aaccaggcca gtctccaaag ctctgatct acaaagtttc caaccgattt	180
tctgggggcc cagacagggt cagtggcagt ggatcaggga cagatttcac actcaagatc	240
aacagagtgg aggctgagga tctgggagtt tattactgct ttcaagggtc acatgttcca	300
ttcacgttcg gctcggggac aaagttggaa ataaaa	336

<210> 459  
 <211> 112  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse C3 Light Chain variable region sequence

<400> 459

Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly
1 5 10 15

Asp Gln Ala Ser Ile Ser Cys Arg Ser Ser Gln Thr Ile Val His Ser
20 25 30

Asn Gly Asn Thr Tyr Leu Glu Trp Tyr Leu Gln Lys Pro Gly Gln Ser
35 40 45

Pro Lys Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser Gly Val Pro
50 55 60

Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile
65 70 75 80

Asn Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys Phe Gln Gly
85 90 95

Ser His Val Pro Phe Thr Phe Gly Ser Gly Thr Lys Leu Glu Ile Lys
100 105 110

<210> 460  
 <211> 69  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse C3 light chain variable framework region 1 (FWR1) sequence

<400> 460

gatgttttga tgacccaaac tccactctcc ctgcctgtca gtcttgagga tcaagcctcc	60
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atctcttgc	69
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<210> 461  
 <211> 23  
 <212> PRT  
 <213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable framework region 1 (FWR1) sequence

<400> 461

Asp	Val	Leu	Met	Thr	Gln	Thr	Pro	Leu	Ser	Leu	Pro	Val	Ser	Leu	Gly
1				5				10						15	

Asp	Gln	Ala	Ser	Ile	Ser	Cys
				20		

<210> 462

<211> 48

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 462

agatctagtc agaccattgt acatagtaat ggaaacacct atttagaa 48

<210> 463

<211> 16

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 463

Arg	Ser	Ser	Gln	Thr	Ile	Val	His	Ser	Asn	Gly	Asn	Thr	Tyr	Leu	Glu
1				5					10					15	

<210> 464

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable framework region 2 (FWR2) sequence

<400> 464

tggtagctgc agaaaccagg ccagtctcca aagctcctga tctac 45

<210> 465

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable framework region 2 (FWR2) sequence

<400> 465

Trp	Tyr	Leu	Gln	Lys	Pro	Gly	Gln	Ser	Pro	Lys	Leu	Leu	Ile	Tyr
1				5				10						15

<210> 466

<211> 21

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 466

aaagtttcca accgattttc t

21

<210> 467

<211> 7

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 467

Lys Val Ser Asn Arg Phe Ser

1 5

<210> 468

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable framework region 3 (FWR3) sequence

<400> 468

ggggtcccag acaggttcag tggcagtgga tcaggacag atttcacact caagatcaac

60

agagtggagg ctgaggatct gggagtttat tactgc

96

<210> 469

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable framework region 3 (FWR3) sequence

<400> 469

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr

1 5 10 15

Leu Lys Ile Asn Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys

20 25 30

<210> 470

<211> 27

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C3 light chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 470

tttcaaggtt cacatgttcc attcacg

27

<210> 471  
<211> 9  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse C3 light chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 471

Phe Gln Gly Ser His Val Pro Phe Thr  
1 5

<210> 472

<400> 472  
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<210> 473

<400> 473  
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<210> 474

<400> 474  
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<210> 475

<400> 475  
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<210> 484

<400> 484  
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<210> 485

<400> 485  
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<210> 486  
<211> 342  
<212> DNA  
<213> Artificial Sequence

<220>

<223> Humanized C3 light chain variable region sequence from IGKV229\*03

<400> 486  
gatattgtga tgacccagac tccactctct ctgtccgtca cccctggaca gccggcctcc 60  
atctcctgca ggtctagtca gaccattgtc catagtaatg gaaacaccta tttggagtgg 120  
tacctgcaga agccaggcca gtctccacag ctctgatct ataaggtttc caaccggttc 180  
tctggagtgc cagatagggtt cagtggcagc gggtcaggga cagatttcac actgaaaatc 240  
agccgggtgg aggctgagga tgttgggggtt tattactgct tccaaggtag ccacgtgcct 300  
ttcaccttcg gcggagggac caaggtggag atcaaacgaa ct 342

<210> 487  
<211> 114  
<212> PRT  
<213> Artificial Sequence

<220>

<223> Humanized C3 light chain variable region sequence from IGKV229\*03

<400> 487

Asp Ile Val Met Thr Gln Thr Pro Leu Ser Leu Ser Val Thr Pro Gly  
1 5 10 15

Gln Pro Ala Ser Ile Ser Cys Arg Ser Ser Gln Thr Ile Val His Ser  
20 25 30

Asn Gly Asn Thr Tyr Leu Glu Trp Tyr Leu Gln Lys Pro Gly Gln Ser  
35 40 45

Pro Gln Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser Gly Val Pro  
50 55 60

Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile  
65 70 75 80

Ser Arg Val Glu Ala Glu Asp Val Gly Val Tyr Tyr Cys Phe Gln Gly  
85 90 95

Ser His Val Pro Phe Thr Phe Gly Gly Thr Lys Val Glu Ile Lys  
100 105 110

Arg Thr

<210> 488  
<211> 69  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 light chain variable framework region 1 (FWR1) acid  
sequence

<400> 488  
gatattgtga tgaccagac tccactctct ctgtccgtca cccctggaca gccggcctcc 60  
atctcctgc 69

<210> 489  
<211> 23  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 light chain variable framework region 1 (FWR1) acid  
sequence

<400> 489

Asp Ile Val Met Thr Gln Thr Pro Leu Ser Leu Ser Val Thr Pro Gly  
1 5 10 15

Gln Pro Ala Ser Ile Ser Cys  
20

<210> 490  
<211> 47  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 light chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 490  
ggtctagtca gaccattgtc catagtaatg gaaacaccta tttggag 47

<210> 491  
<211> 16  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 light chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 491



Arg Ser Ser Gln Thr Ile Val His Ser Asn Gly Asn Thr Tyr Leu Glu  
1 5 10 15

<210> 492  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 light chain variable framework region 2 (FWR2) acid  
sequence

<400> 492  
tggtagctgc agaagccagg ccagttctca cagctcctga tctat 45

<210> 493  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 light chain variable framework region 2 (FWR2) acid  
sequence

<400> 493

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Gln Leu Leu Ile Tyr  
1 5 10 15

<210> 494  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 light chain variable complementarity determining  
regions 2 (CDR2) sequence

<400> 494  
aaggtttcca accggttctc t 21

<210> 495  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 light chain variable complementarity determining  
regions 2 (CDR2) sequence

<400> 495

Lys Val Ser Asn Arg Phe Ser  
1 5

<210> 496  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 light chain variable framework region 3 (FWR3) acid  
sequence

<400> 496  
 ggagtgccag ataggttcag tggcagcggg tcaggacag atttcacact gaaaatcagc 60  
 cgggtggagg ctgaggatgt tggggtttat tactgc 96

<210> 497  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Humanized C3 light chain variable framework region 3 (FWR3) acid  
 sequence

<400> 497  
 Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Val Gly Val Tyr Tyr Cys  
 20 25 30

<210> 498  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Humanized C3 light chain variable complementarity determining  
 regions 3 (CDR3) sequence

<400> 498  
 ttccaaggta gccacgtgcc tttcacc 27

<210> 499  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Humanized C3 light chain variable complementarity determining  
 regions 3 (CDR3) sequence

<400> 499  
 Phe Gln Gly Ser His Val Pro Phe Thr  
 1 5

<210> 500  
 <211> 666  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Humanized C3 lambda light chain sequence

<400> 500  
 gatattgtga tgaccagac tccactctct ctgtccgtca cccctggaca gccggcctcc 60  
 atctcctgca ggtctagtca gaccattgtc catagtaatg gaaacaccta tttggagtgg 120  
 tacctgcaga agccaggcca gtctccacag ctctgatctc ataaggtttc caaccggttc 180



Ser Pro Val Lys Ala Gly Val Glu Thr Thr Thr Pro Ser Lys Gln Ser  
165 170 175

Asn Asn Lys Tyr Ala Ala Ser Ser Tyr Leu Ser Leu Thr Pro Glu Gln  
180 185 190

Trp Lys Ser His Arg Ser Tyr Ser Cys Gln Val Thr His Glu Gly Ser  
195 200 205

Thr Val Glu Lys Thr Val Ala Pro Thr Glu Cys Ser  
210 215 220

<210> 502  
<211> 666  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C3 Kappa light chain

<400> 502  
gatattgtga tgaccagac tccactctct ctgtccgtca cccctggaca gccggcctcc 60  
atctcctgca ggtctagtca gaccattgtc catagtaatg gaaacaccta tttggagtgg 120  
tacctgcaga agccaggcca gtctccacag ctctgatct ataaggtttc caaccggttc 180  
tctggagtgc cagatagggtt cagtggcagc gggtcaggga cagatttcac actgaaaatc 240  
agccgggtgg aggctgagga tgttgggggtt tattactgct tccaaggtag ccacgtgcct 300  
ttcaccttcg gcggaggagc caaggtggag atcaaacgaa ctacgggtggc tgcaccatct 360  
gtcttcatct tcccgccatc tgatgagcag ttgaaatctg gaactgcctc tgttgtgtgc 420  
ctgctgaata acttctatcc cagagaggcc aaagtacagt ggaagggtgga taacgccctc 480  
caatcgggta actcccagga gagtgtcaca gagcaggaca gcaaggacag cacctacagc 540  
ctcagcagca cctgacgct gagcaaagca gactacgaga aacacaaagt ctacgcctgc 600  
gaagtcaccc atcagggcct gagctcgccc gtcacaaaga gcttcaacag gggagagtgt 660  
tagtaa 666

<210> 503  
<211> 220  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C3 Kappa light chain

<400> 503

Asp Ile Val Met Thr Gln Thr Pro Leu Ser Leu Ser Val Thr Pro Gly  
1 5 10 15

Gln Pro Ala Ser Ile Ser Cys Arg Ser Ser Gln Thr Ile Val His Ser  
20 25 30

Asn Gly Asn Thr Tyr Leu Glu Trp Tyr Leu Gln Lys Pro Gly Gln Ser

35

40

45

Pro Gln Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser Gly Val Pro  
50 55 60

Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile  
65 70 75 80

Ser Arg Val Glu Ala Glu Asp Val Gly Val Tyr Tyr Cys Phe Gln Gly  
85 90 95

Ser His Val Pro Phe Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys  
100 105 110

Arg Thr Thr Val Ala Ala Pro Ser Val Phe Ile Phe Pro Pro Ser Asp  
115 120 125

Glu Gln Leu Lys Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn  
130 135 140

Phe Tyr Pro Arg Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu  
145 150 155 160

Gln Ser Gly Asn Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp  
165 170 175

Ser Thr Tyr Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr  
180 185 190

Glu Lys His Lys Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser  
195 200 205

Ser Pro Val Thr Lys Ser Phe Asn Arg Gly Glu Cys  
210 215 220

<210> 504

<400> 504  
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<210> 505

<211> 321

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse C8 heavy chain variable region sequence

<400> 505

gaagtgatgg tcgtggaaag cggcgggtggt ctggtaaagc cggggggatc ccttaagctt 60

tcttgcgccg catccgggtt cacgttctcc ggctatgcc a tgctctgggt cgcacagact 120

cccgaaaagc gcttggaaatg ggtggccact atctcctccg gggggacgta catctactac 180

cccgcagctg tgaaggaag atttacaata tctcgcgaca acgcaaaaaa taccttgat 240

cttcaaatga gctccctgcg gtcagaggac actgccatgt actattgcgc ccgcctgggc 300  
ggcgacaatt actatgagta t 321

<210> 506  
<211> 107  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse C8 heavy chain variable region sequence

<400> 506

Glu Val Met Val Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
20 25 30

Ala Met Ser Trp Val Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val  
35 40 45

Ala Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr  
65 70 75 80

Leu Gln Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Met Tyr Tyr Cys  
85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr  
100 105

<210> 507  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse C8 heavy chain variable complementarity determining region  
1 (CDR1) sequence

<400> 507  
ggctatgcc tgtcc 15

<210> 508  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse C8 heavy chain variable complementarity determining region  
1 (CDR1) sequence

<400> 508

Gly Tyr Ala Met Ser  
1 5

<210> 509  
 <211> 51  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse C8 heavy chain variable complementarity determining region  
 2 (CDR2) sequence  
  
 <400> 509  
 actatctcct ccggggggac gtacatctac taccccgaca gtgtgaaagg a 51

<210> 510  
 <211> 17  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse C8 heavy chain variable complementarity determining region  
 2 (CDR2) sequence  
  
 <400> 510  
  
 Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys  
 1 5 10 15

Gly

<210> 511  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse C8 heavy chain variable complementarity determining region  
 3 (CDR3) sequence  
  
 <400> 511  
 ctgggcggcg acaattacta tgagtat 27

<210> 512  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse C8 heavy chain variable complementarity determining region  
 3 (CDR3) sequence  
  
 <400> 512  
  
 Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr  
 1 5

<210> 513  
  
 <400> 513  
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<400> 514  
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<210> 525

<211> 354

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable region sequence from IGHV321\*04

<400> 525

gaggtgcagc tggaggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60

tcctgtgcag cctctggatt caccttcagt ggctatgccca tgagctgggt ccgccaggct 120

ccagggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac 180

cctgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240



ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagactgggc 300

ggcgataact attatgaata ttggggcaaa gggaccacgg tcaccgtctc ctcc 354

<210> 526

<211> 118

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable region sequence from IGHV321\*04

<400> 526

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
20 25 30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Trp Gly Lys Gly Thr  
100 105 110

Thr Val Thr Val Ser Ser  
115

<210> 527

<211> 90

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable framework region 1 (FWR1)  
sequence

<400> 527

gaggtgcagc tgggtggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60

tcctgtgcag cctctggatt caccttcagt 90

<210> 528

<211> 30

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable framework region 1 (FWR1)  
sequence

<400> 528

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 529

<211> 15

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable complementarity determining  
region 1 (CDR1) sequence

<400> 529

ggctatgcc ttagc

15

<210> 530

<211> 5

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable complementarity determining  
region 1 (CDR1) sequence

<400> 530

Gly Tyr Ala Met Ser  
1 5

<210> 531

<211> 42

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable framework region 2 (FWR2)  
sequence

<400> 531

tgggtccgcc aggtccagg gaaggggctg gagtgggtct ca

42

<210> 532

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable framework region 2 (FWR2)  
sequence

<400> 532

Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val Ser  
1 5 10

<210> 533  
 <211> 51  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized C8 heavy chain variable complementarity determining region 2 (CDR2) sequence  
  
 <400> 533  
 accattagta gtggcggaac ctacatatac taccctgact cagtgaaggg c 51

<210> 534  
 <211> 17  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized C8 heavy chain variable complementarity determining region 2 (CDR2) sequence

<400> 534  
  
 Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys  
 1 5 10 15

Gly

<210> 535  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized C8 heavy chain variable framework region 3 (FWR3) sequence  
  
 <400> 535  
 cgattcacca tctccagaga caacgccaaag aactcactgt atctgcaaat gaacagcctg 60  
 agagccgagg acacggccgt gtattactgt gcgaga 96

<210> 536  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Humanized C8 heavy chain variable framework region 3 (FWR3) sequence

<400> 536  
  
 Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr Leu Gln  
 1 5 10 15

Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys Ala Arg  
 20 25 30

<210> 537  
 <211> 27  
 <212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable complementarity determining region 3 (CDR3) sequence

<400> 537

ctgggcggcg ataactatta tgaatat

27

<210> 538

<211> 9

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C8 heavy chain variable complementarity determining region 3 (CDR3) sequence

<400> 538

Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr

1

5

<210> 539

<211> 1350

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C8 IgG1 heavy chain sequence

<400> 539

gaggtgcagc tgggtggagtc tggggggaggc ctggtcaagc ctgggggggtc cctgagactc 60

tcctgtgcag cctctggatt caccttcagt ggctatgcc a tgagctgggt ccgccaggct 120

ccagggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac 180

cctgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240

ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagactgggc 300

ggcgataact attatgaata ttggggcaaa gggaccacgg tcaccgtctc ctccgctagc 360

accaagggcc catcggtctt ccccttgga ccctcctcca agagcacctc tgggggcaca 420

gcggccctgg gctgcctggt caaggactac ttccccgaac cggtgacggt gtcgtggaac 480

tcaggcgccc tgaccagcgg cgtgcacacc ttcccggtg tcctacagtc ctcaggactc 540

tactccctca gcagcgtggt gacagtgtcc tccagcagct tgggcaccca gacctacatc 600

tgcaacgtga atcacaagcc cagcaacacc aaggtggaca agaaagttga gcccaaactc 660

tgtgacaaaa ctacacatg cccaccgtgc ccagcacctg aactcctggg gggaccgtca 720

gtcttctctt tcccccaaa acccaaggac accctcatga tctccggac ccctgaggtc 780

acatgcgtgg tgggtggacgt gagccacgaa gaccctgagg tcaagttcaa ctggtacgtg 840

gacggcgtgg aggtgcataa tgccaagaca aagccgcggg aggagcagta caacagcacg 900

taccgtgtgg tcagcgtcct caccgtcctg caccaggact ggctgaatgg caaggagtac 960

aagtgcagg tctccaacaa agccctccca gccccatcg agaaaacat ctccaaagcc 1020

aaagggcagc cccgagaacc acaggtgtac accctgcccc catcccggga ggagatgacc 1080

aagaaccagg tcagcctgac ctgcctggtc aaaggcttct atcccagcga catcgccgtg	1140
gagtgggaga gcaatgggca gccggagaac aactacaaga ccacgcctcc cgtgctggac	1200
tccgacggct cttctttcct ctacagcaag ctcaccgtgg acaagagcag gtggcagcag	1260
gggaacgtct tctcatgctc cgtgatgcat gaggctctgc acaaccacta cacgcagaag	1320
agcctctccc tgtctccggg taaatgataa	1350

<210> 540  
 <211> 448  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Humanized C8 IgG1 heavy chain sequence

<400> 540

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly	
1                    5                    10                    15	

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr	
20                    25                    30	

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val	
35                    40                    45	

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val	
50                    55                    60	

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr	
65                    70                    75                    80	

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys	
85                    90                    95	

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Trp Gly Lys Gly Thr	
100                    105                    110	

Thr Val Thr Val Ser Ser Ala Ser Thr Lys Gly Pro Ser Val Phe Pro	
115                    120                    125	

Leu Ala Pro Ser Ser Lys Ser Thr Ser Gly Gly Thr Ala Ala Leu Gly	
130                    135                    140	

Cys Leu Val Lys Asp Tyr Phe Pro Glu Pro Val Thr Val Ser Trp Asn	
145                    150                    155                    160	

Ser Gly Ala Leu Thr Ser Gly Val His Thr Phe Pro Ala Val Leu Gln	
165                    170                    175	

Ser Ser Gly Leu Tyr Ser Leu Ser Ser Val Val Thr Val Pro Ser Ser	
180                    185                    190	

Ser Leu Gly Thr Gln Thr Tyr Ile Cys Asn Val Asn His Lys Pro Ser  
195 200 205

Asn Thr Lys Val Asp Lys Lys Val Glu Pro Lys Ser Cys Asp Lys Thr  
210 215 220

His Thr Cys Pro Pro Cys Pro Ala Pro Glu Leu Leu Gly Gly Pro Ser  
225 230 235 240

Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser Arg  
245 250 255

Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser His Glu Asp Pro  
260 265 270

Glu Val Lys Phe Asn Trp Tyr Val Asp Gly Val Glu Val His Asn Ala  
275 280 285

Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn Ser Thr Tyr Arg Val Val  
290 295 300

Ser Val Leu Thr Val Leu His Gln Asp Trp Leu Asn Gly Lys Glu Tyr  
305 310 315 320

Lys Cys Lys Val Ser Asn Lys Ala Leu Pro Ala Pro Ile Glu Lys Thr  
325 330 335

Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu  
340 345 350

Pro Pro Ser Arg Glu Glu Met Thr Lys Asn Gln Val Ser Leu Thr Cys  
355 360 365

Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser  
370 375 380

Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro Pro Val Leu Asp  
385 390 395 400

Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser  
405 410 415

Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala  
420 425 430

Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys  
435 440 445

<210> 541

<211> 1338

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C8 IgG2 heavy chain sequence

<400> 541

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gagggtgcagc tgggtggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc      60
tcctgtgcag cctctggatt cacccttcagt ggctatgcca tgagctgggt ccgccaggct      120
ccaggggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac      180
cctgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat      240
ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagactgggc      300
ggcgataact attatgaata ttggggcaaa gggaccacgg tcaccgtctc ctccgcctcc      360
accaagggcc catcgggtctt ccccttggcg ccctgtctca ggagcacctc cgagagcaca      420
gccgccctgg gctgcctggg caaggactac ttccccgaac cggtgacggg gtcgtggaac      480
tcaggcgctc tgaccagcgg cgtgcacacc ttcccagctg tcctacagtc ctcaggactc      540
tactccctca gcagcgtggg gaccgtgccc tccagcaact tcggcaccca gacctacacc      600
tgcaacgtag atcacaagcc cagcaacacc aagggtggaca agacagttga gcgcaaatgt      660
tgtgtcgagt gcccaccgtg cccagcacca cctgtggcag gaccgtcagt cttcctcttc      720
ccccaaaac ccaaggacac cctcatgac tcccggacc ctgaggtcac gtgcgtggtg      780
gtggacgtga gccacgaaga ccccgaggtc cagttcaact ggtacgtgga cggcgtggag      840
gtgcataatg ccaagacaaa gccacgggag gagcagttca acagcacgtt ccgtgtggtc      900
agcgtcctca ccgttgtgca ccaggactgg ctgaacggca aggagtacaa gtgcaaggtc      960
tccaacaaag gcctcccagc ccccatcgag aaaaccatct ccaaaaccaa agggcagccc     1020
cgagaaccac aggtgtacac cctgccccca tcccgggagg agatgaccaa gaaccaggtc     1080
agcctgacct gcctgggtcaa aggcttctac cccagcgaca tcgccgtgga gtgggagagc     1140
aatgggcagc cggagaacaa ctacaagacc acacctccca tgctggactc cgacggctcc     1200
ttcttctctt acagcaagct caccgtggac aagagcaggt ggcagcaggg gaacgtcttc     1260
tcatgctccg tgatgcatga ggctctgcac aaccactaca cgcagaagag cctctccctg     1320
tctccgggta aatagtaa                                     1338
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<210> 542

<211> 444

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C8 IgG2 heavy chain sequence

<400> 542

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Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly
1          5          10         15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr
20         25         30
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Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Trp Gly Lys Gly Thr  
100 105 110

Thr Val Thr Val Ser Ser Ala Ser Thr Lys Gly Pro Ser Val Phe Pro  
115 120 125

Leu Ala Pro Cys Ser Arg Ser Thr Ser Glu Ser Thr Ala Ala Leu Gly  
130 135 140

Cys Leu Val Lys Asp Tyr Phe Pro Glu Pro Val Thr Val Ser Trp Asn  
145 150 155 160

Ser Gly Ala Leu Thr Ser Gly Val His Thr Phe Pro Ala Val Leu Gln  
165 170 175

Ser Ser Gly Leu Tyr Ser Leu Ser Ser Val Val Thr Val Pro Ser Ser  
180 185 190

Asn Phe Gly Thr Gln Thr Tyr Thr Cys Asn Val Asp His Lys Pro Ser  
195 200 205

Asn Thr Lys Val Asp Lys Thr Val Glu Arg Lys Cys Cys Val Glu Cys  
210 215 220

Pro Pro Cys Pro Ala Pro Pro Val Ala Gly Pro Ser Val Phe Leu Phe  
225 230 235 240

Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val  
245 250 255

Thr Cys Val Val Val Asp Val Ser His Glu Asp Pro Glu Val Gln Phe  
260 265 270

Asn Trp Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro  
275 280 285

Arg Glu Glu Gln Phe Asn Ser Thr Phe Arg Val Val Ser Val Leu Thr  
290 295 300



Val Val His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val  
305 310 315 320

Ser Asn Lys Gly Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Thr  
325 330 335

Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Arg  
340 345 350

Glu Glu Met Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly  
355 360 365

Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro  
370 375 380

Glu Asn Asn Tyr Lys Thr Thr Pro Pro Met Leu Asp Ser Asp Gly Ser  
385 390 395 400

Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln Gln  
405 410 415

Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His  
420 425 430

Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys  
435 440

<210> 543  
<211> 306  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse C8 light chain variable region sequence

<400> 543  
gacatcgtca ttacgcagac ccctgccagt cttgccgttt ctctgggcca gagggccact 60  
atcagttaca gggcgagtaa gtctgtgagt accagcggct atagttacat gcattggaac 120  
cagcagaaac cgggacagcc accacgcctg cttatttatc tgggtgtctaa tcttgagtcc 180  
gggggtgcccc ccaggttcag cggcagcggc tctgggaccg acttcacact caacattcat 240  
ccagtggaag aagaggacgc tgctacatac tactgtcaac acattcggga actgaccagg 300  
agtgaa 306

<210> 544  
<211> 102  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse C8 light chain variable region sequence

<400> 544

Asp Ile Val Ile Thr Gln Thr Pro Ala Ser Leu Ala Val Ser Leu Gly

1	5	10	15
Gln Arg Ala Thr Ile Ser Tyr Arg Ala Ser Lys Ser Val Ser Thr Ser			
20	25	30	
Gly Tyr Ser Tyr Met His Trp Asn Gln Gln Lys Pro Gly Gln Pro Pro			
35	40	45	
Arg Leu Leu Ile Tyr Leu Val Ser Asn Leu Glu Ser Gly Val Pro Ala			
50	55	60	
Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Asn Ile His			
65	70	75	80
Pro Val Glu Glu Glu Asp Ala Ala Thr Tyr Tyr Cys Gln His Ile Arg			
85	90	95	
Glu Leu Thr Arg Ser Glu			
100			

<210> 545  
 <211> 45  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse C8 light chain variable complementarity determining region  
 1 (CDR1) sequence

<400> 545  
 agggcgagta agtctgtgag taccagcggc tatagttaca tgcac 45

<210> 546  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse C8 light chain variable complementarity determining region  
 1 (CDR1) sequence

<400> 546

Arg Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser Tyr Met His
1 5 10 15

<210> 547  
 <211> 21  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse C8 light chain variable complementarity determining region  
 2 (CDR2) sequence

<400> 547  
 ctggtgtcta atcttgagtc c 21

<210> 548

<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse C8 light chain variable complementarity determining region  
2 (CDR2) sequence

<400> 548

Leu Val Ser Asn Leu Glu Ser  
1 5

<210> 549  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse C8 light chain variable complementarity determining region  
3 (CDR3) sequence

<400> 549  
caacacattc gggaactgac caggagtgaa 30

<210> 550  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse C8 light chain variable complementarity determining region  
3 (CDR3) sequence

<400> 550

Gln His Ile Arg Glu Leu Thr Arg Ser Glu  
1 5 10

<210> 551

<400> 551  
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<400> 564  
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<210> 565

<211> 342

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C8 light chain variable region sequence from NCBI  
germline z00023

<400> 565  
gacatcgtga tgacccagtc tccagactcc ctggctgtgt ctctgggcga gagggccacc 60  
atcaactgca gggccagcaa gagtgttagc accagcggct acagctacat gactggtac 120  
cagcagaaac caggacagcc tcctaagctg ctcatctacc tgggtgtctaa cctggaatcc 180  
gggggtccctg accgattcag tggcagcggg tctgggacag atttcactct caccatcagc 240  
agcctgcagg ctgaagatgt ggcagtttat tactgtcaac acattcggga actgaccagg 300  
agtgaattcg gcggagggac caaggtggag atcaaacgaa ct 342

<210> 566

<211> 114

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C8 light chain variable region sequence from NCBI  
germline z00023

<400> 566

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu Ala Val Ser Leu Gly  
1 5 10 15

Glu Arg Ala Thr Ile Asn Cys Arg Ala Ser Lys Ser Val Ser Thr Ser  
20 25 30

Gly Tyr Ser Tyr Met His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro  
35 40 45

Lys Leu Leu Ile Tyr Leu Val Ser Asn Leu Glu Ser Gly Val Pro Asp  
50 55 60

Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser  
65 70 75 80

Ser Leu Gln Ala Glu Asp Val Ala Val Tyr Tyr Cys Gln His Ile Arg  
85 90 95

Glu Leu Thr Arg Ser Glu Phe Gly Gly Gly Thr Lys Val Glu Ile Lys  
100 105 110

Arg Thr

<210> 567

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Humanized C8 light chain variable framework region 1 (FWR1)  
sequence

<400> 567

gacatcgtga tgaccagtc tccagactcc ctggctgtgt ctctgggcga gagggccacc 60

atcaactgc 69

<210> 568

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Humanized C8 light chain variable framework region 1 (FWR1)  
sequence

<400> 568

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu Ala Val Ser Leu Gly  
1 5 10 15

Glu Arg Ala Thr Ile Asn Cys  
20

<210> 569  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C8 light chain variable complementarity determining  
region 1 (CDR1) sequence

<400> 569  
agggccagca agagtgttag caccagcggc tacagctaca tg 42

<210> 570  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C8 light chain variable complementarity determining  
region 1 (CDR1) sequence

<400> 570

Arg Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser Tyr Met  
1 5 10

<210> 571  
<211> 48  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C8 light chain variable framework region 2 (FWR2)  
sequence

<400> 571  
cactggtacc agcagaaacc aggacagcct cctaagctgc tcatttac 48

<210> 572  
<211> 16  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C8 light chain variable framework region 2 (FWR2)  
sequence

<400> 572

His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 573  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C8 light chain variable complementarity determining  
region 2 (CDR2) sequence

<400> 573  
ctggtgtcta acctggaatc c 21

<210> 574  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C8 light chain variable complementarity determining  
region 2 (CDR2) sequence

<400> 574  
  
Leu Val Ser Asn Leu Glu Ser  
1 5

<210> 575  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C8 light chain variable framework region 3 (FWR3)  
sequence

<400> 575  
ggggtccctg accgattcag tggcagcggg tctgggacag atttcactct caccatcagc 60  
  
agcctgcagg ctgaagatgt ggcagtttat tactgt 96

<210> 576  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C8 light chain variable framework region 3 (FWR3)  
sequence

<400> 576  
  
Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Thr Ile Ser Ser Leu Gln Ala Glu Asp Val Ala Val Tyr Tyr Cys  
20 25 30

<210> 577  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C8 light chain variable complementarity determining  
region 3 (CDR3) sequence

<400> 577  
caacacattc gggaactgac caggagtga 30

<210> 578  
<211> 10

<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Humanized C8 light chain variable complementarity determining region 3 (CDR3) sequence

<400> 578

Gln His Ile Arg Glu Leu Thr Arg Ser Glu  
1 5 10

<210> 579  
<211> 666  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Humanized C8 Lambda light chain sequence

<400> 579  
gacatcgtga tgaccagtc tccagactcc ctggctgtgt ctctgggcga gagggccacc 60  
atcaactgca gggccagcaa gagtgttagc accagcggct acagctacat gactggtac 120  
cagcagaaac caggacagcc tcctaagctg ctcatctacc tgggtgtctaa cctggaatcc 180  
gggggtccctg accgattcag tggcagcggg tctgggacag atttcactct caccatcagc 240  
agcctgcagg ctgaagatgt ggcagtttat tactgtcaac acattcggga actgaccagg 300  
agtgaattcg gcggaggggac caaggtggag atcaaacgaa ctggtcagcc caaggctgcc 360  
ccctcgggtca ctctgttccc gccctcctct gaggagcttc aagccaacaa ggccacactg 420  
gtgtgtctca taagtgaact ctaccggga gccgtgacag tggcctggaa ggcagatagc 480  
agccccgtca aggcgggagt ggagaccacc acaccctcca aacaaagcaa caacaagtac 540  
gcggccagca gctatctgag cctgacgcct gagcagtggga agtcccacag aagctacagc 600  
tgccagggtca cgcataaagg gagcaccgtg gagaagacag tggcccctac agaattgtca 660  
tagtaa 666

<210> 580  
<211> 220  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Humanized C8 Lambda light chain sequence

<400> 580

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu Ala Val Ser Leu Gly  
1 5 10 15

Glu Arg Ala Thr Ile Asn Cys Arg Ala Ser Lys Ser Val Ser Thr Ser  
20 25 30

Gly Tyr Ser Tyr Met His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro  
35 40 45



Lys Leu Leu Ile Tyr Leu Val Ser Asn Leu Glu Ser Gly Val Pro Asp  
 50 55 60  
 Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser  
 65 70 75 80  
 Ser Leu Gln Ala Glu Asp Val Ala Val Tyr Tyr Cys Gln His Ile Arg  
 85 90 95  
 Glu Leu Thr Arg Ser Glu Phe Gly Gly Gly Thr Lys Val Glu Ile Lys  
 100 105 110  
 Arg Thr Gly Gln Pro Lys Ala Ala Pro Ser Val Thr Leu Phe Pro Pro  
 115 120 125  
 Ser Ser Glu Glu Leu Gln Ala Asn Lys Ala Thr Leu Val Cys Leu Ile  
 130 135 140  
 Ser Asp Phe Tyr Pro Gly Ala Val Thr Val Ala Trp Lys Ala Asp Ser  
 145 150 155 160  
 Ser Pro Val Lys Ala Gly Val Glu Thr Thr Thr Pro Ser Lys Gln Ser  
 165 170 175  
 Asn Asn Lys Tyr Ala Ala Ser Ser Tyr Leu Ser Leu Thr Pro Glu Gln  
 180 185 190  
 Trp Lys Ser His Arg Ser Tyr Ser Cys Gln Val Thr His Glu Gly Ser  
 195 200 205  
 Thr Val Glu Lys Thr Val Ala Pro Thr Glu Cys Ser  
 210 215 220

<210> 581  
 <211> 666  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Humanized C8 Kappa light chain sequence

<400> 581  
 gacatcgtga tgaccagtc tccagactcc ctggctgtgt ctctgggcga gagggccacc 60  
 atcaactgca gggccagcaa gagggttagc accagcggct acagctacat gactgggtac 120  
 cagcagaaac caggacagcc tcctaagctg ctcatctacc tgggtgtctaa cctggaatcc 180  
 ggggtccctg accgattcag tggcagcggg tctgggacag atttcactct caccatcagc 240  
 agcctgcagg ctgaagatgt ggcagtttat tactgtcaac acattcgga actgaccagg 300  
 agtgaattcg gcggaggag caaggtggag atcaaacgaa ctacggtggc tgcaccatct 360  
 gtcttcatct tcccgccatc tgatgagcag ttgaaatctg gaactgcctc tgttgtgtgc 420  
 ctgctgaata acttctatcc cagagaggcc aaagtacagt ggaaggtgga taacgccctc 480

caatcgggta actcccagga gagtgtcaca gagcaggaca gcaaggacag cacctacagc	540
ctcagcagca ccctgacgct gagcaaagca gactacgaga aacacaaagt ctacgcctgc	600
gaagtcaccc atcagggcct gagctcgccc gtcacaaaga gcttcaacag gggagagtgt	660
tagtaa	666

<210> 582  
 <211> 220  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Humanized C8 Kappa light chain sequence

<400> 582

Asp	Ile	Val	Met	Thr	Gln	Ser	Pro	Asp	Ser	Leu	Ala	Val	Ser	Leu	Gly
1				5					10					15	

Glu	Arg	Ala	Thr	Ile	Asn	Cys	Arg	Ala	Ser	Lys	Ser	Val	Ser	Thr	Ser
			20					25					30		

Gly	Tyr	Ser	Tyr	Met	His	Trp	Tyr	Gln	Gln	Lys	Pro	Gly	Gln	Pro	Pro
		35					40					45			

Lys	Leu	Leu	Ile	Tyr	Leu	Val	Ser	Asn	Leu	Glu	Ser	Gly	Val	Pro	Asp
	50					55					60				

Arg	Phe	Ser	Gly	Ser	Gly	Ser	Gly	Thr	Asp	Phe	Thr	Leu	Thr	Ile	Ser
65					70					75					80

Ser	Leu	Gln	Ala	Glu	Asp	Val	Ala	Val	Tyr	Tyr	Cys	Gln	His	Ile	Arg
				85					90					95	

Glu	Leu	Thr	Arg	Ser	Glu	Phe	Gly	Gly	Gly	Thr	Lys	Val	Glu	Ile	Lys
			100					105					110		

Arg	Thr	Thr	Val	Ala	Ala	Pro	Ser	Val	Phe	Ile	Phe	Pro	Pro	Ser	Asp
			115					120					125		

Glu	Gln	Leu	Lys	Ser	Gly	Thr	Ala	Ser	Val	Val	Cys	Leu	Leu	Asn	Asn
	130					135					140				

Phe	Tyr	Pro	Arg	Glu	Ala	Lys	Val	Gln	Trp	Lys	Val	Asp	Asn	Ala	Leu
145					150					155					160

Gln	Ser	Gly	Asn	Ser	Gln	Glu	Ser	Val	Thr	Glu	Gln	Asp	Ser	Lys	Asp
				165					170					175	

Ser	Thr	Tyr	Ser	Leu	Ser	Ser	Thr	Leu	Thr	Leu	Ser	Lys	Ala	Asp	Tyr
			180					185					190		

Glu	Lys	His	Lys	Val	Tyr	Ala	Cys	Glu	Val	Thr	His	Gln	Gly	Leu	Ser
		195					200					205			

Ser Pro Val Thr Lys Ser Phe Asn Arg Gly Glu Cys  
 210 215 220

<210> 583

<400> 583  
 000

<210> 584  
 <211> 942  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CART E6 CD8 sequence

<400> 584  
 gaggtccagc tggttgagag tggcgggtggg ctggttaagc ctggcggctc cctgcggtctg 60  
 agctgcgccg cgagtggatt tactttcagc cgatatggga tgagttgggt gcggcaagct 120  
 cccgggaaga ggctggaatg ggtctcaaca atctccgggg ggggcactta catctattac 180  
 cccgactcag tcaaggggag atttaccatt tcacgagaca acgctaagaa taccctgtat 240  
 ttgcagatga attctctgag agcagaggac acagctgttt actattgtac ccgcgacaac 300  
 tatggcagga actacgacta cggtatggac tattggggac aaggacatt ggttacagtg 360  
 agcagtggcg gcggggggcag cggaggagga ggcagcgggtg gggggggcag cgagatagtg 420  
 ctcacgcagt caccgcgcac tctcagtctc tcacctgggg aacgagctac cctgacgtgc 480  
 tctgctacct cctcagtgtc atatattcac tggtatcagc aacggcccgg gcagtcccct 540  
 agattgctca tttatagtag ctctaacttg gcctcaggta tccctgcacg attttctgga 600  
 tctggttcag gttctgatta caccctcact atctctagcc tggagcctga agactttgcc 660  
 gtttattact gccagcagag gtctagctcc ccattcacct ttgggagtgg gaccaagggt 720  
 gaaattaaaa cgacaacccc ggccccccaga ccaccaacgc cagccccac catcgccagc 780  
 caaccctgt ctctgagacc agaagcctgt aggcctgccg ccggtggagc tgtgcacaca 840  
 agaggactgg atttcgcctg tgatatctac atttggggcc cgctcgcagg cacatgtgga 900  
 gtgctcctcc tctccctggg gattaccctg tactgctgat aa 942

<210> 585  
 <211> 312  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> CART E6 CD8 sequence

<400> 585

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
 1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Arg Tyr  
 20 25 30

Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys Arg Leu Glu Trp Val  
 35 40 45

Ser Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
 50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr  
 65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
 85 90 95

Thr Arg Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr Gly Met Asp Tyr Trp  
 100 105 110

Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly  
 115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Ile Val Leu Thr Gln Ser  
 130 135 140

Pro Ala Thr Leu Ser Leu Ser Pro Gly Glu Arg Ala Thr Leu Thr Cys  
 145 150 155 160

Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp Tyr Gln Gln Arg Pro  
 165 170 175

Gly Gln Ser Pro Arg Leu Leu Ile Tyr Ser Thr Ser Asn Leu Ala Ser  
 180 185 190

Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Ser Asp Tyr Thr  
 195 200 205

Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys  
 210 215 220

Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly Ser Gly Thr Lys Val  
 225 230 235 240

Glu Ile Lys Thr Thr Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro  
 245 250 255

Thr Ile Ala Ser Gln Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro  
 260 265 270

Ala Ala Gly Gly Ala Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp  
 275 280 285

Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu  
 290 295 300

Ser Leu Val Ile Thr Leu Tyr Cys  
305 310

<210> 586  
<211> 960  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CART C2 CD8 CD8 sequence - NCD81shuMNC2scFvCD8ecd fragment CD8  
transmembraneC

<400> 586  
gaagtgcagc tcgtagagag tggcggggga ctggtgaagc ccggtggaag cctcagactc 60  
agttgcgccg cctcaggttt cactttttca ggttacgcca tgtcctgggt aagacaggca 120  
ccggggaaag gactcgagtg ggtgtctact atcagctcag gaggcactta tatatattat 180  
cctgactctg taaaaggccg atttacgatt tctcgcgaca atgcaaagaa ctccctctac 240  
ctccaaatga acagtcttag ggcagaagac actgctgtat actattgtgc acgcctcggc 300  
ggcgacaact actacgagta ctttgacgtg tgggggaaag ggactaccgt gacagtttca 360  
agcggaggag gtggctcagg tggaggcggg tcaggggggg gaggaagtga tattgtgctc 420  
acacaatccc cagcctccct ggctgtgtct cccggccaac gcgctacaat tacatgtcgg 480  
gcctccaaaa gcgtgagcac cagcggctac agctacatgc actggtatca acagaaacca 540  
ggacaacccc ccaaactgtt gatttatctc gcttcaaact tggagtccgg cgtgcctgcg 600  
cgcttttcag ggagtgggag cggcacagat ttacgctga ctatcaacct cgtagaagca 660  
aacgatacag cgaattatta ttgtcaacat tcccgggaac tcccctttac gttcggcggg 720  
ggcacaaagg tcgaaattaa gagaaccacg acaaccccgg ccccgagacc accaacgcca 780  
gccccacca tcgccagcca acccctgtct ctgagaccag aagcctgtag gcctgccgcc 840  
ggtggagctg tgcacacaag aggactggat ttgcctgtg atatctacat ttgggccccg 900  
ctcgcaggca catgtggagt gtcctcctc tccctggtga ttaccctgta ctgctgataa 960

<210> 587  
<211> 318  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CART C2 CD8 CD8 sequence - NCD81shuMNC2scFvCD8ecd fragment CD8  
transmembraneC

<400> 587

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Gly Tyr  
20 25 30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45

Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val  
 50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
 65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
 85 90 95

Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr Phe Asp Val Trp Gly  
 100 105 110

Lys Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly  
 115 120 125

Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Leu Thr Gln Ser Pro  
 130 135 140

Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala Thr Ile Thr Cys Arg  
 145 150 155 160

Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser Tyr Met His Trp Tyr  
 165 170 175

Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Leu Ala Ser  
 180 185 190

Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly  
 195 200 205

Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu Ala Asn Asp Thr Ala  
 210 215 220

Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro Phe Thr Phe Gly Gly  
 225 230 235 240

Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr Thr Pro Ala Pro Arg  
 245 250 255

Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser Leu Arg  
 260 265 270

Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr Arg Gly  
 275 280 285

Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr  
 290 295 300

Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys  
 305 310 315

<210> 588  
<211> 339  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8/41BB sequence - N CD8 transmembrane 41BBC

<400> 588  
acgacaaccc cggccccag accaccaacg ccagcccca ccatgccag ccaaccctg 60  
tctctgagac cagaagcctg taggcctgcc gccggtggag ctgtgcacac aagaggactg 120  
gatttcgcct gtgatatcta catttgggcc ccgctcgag gcacatgtgg agtgctcctc 180  
ctctccctgg tgattaccct gtactgcaaa aggggccgca aaaaactcct ttacattttt 240  
aagcagcctt ttatgaggcc agtacagacg actcaagagg aagacgggtg ctcatgccgc 300  
tttctgagg aggaggaagg aggggtcgaa ctgtgataa 339

<210> 589  
<211> 111  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CD8/41BB sequence - N CD8 transmembrane 41BBC

<400> 589

Thr Thr Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala  
1 5 10 15

Ser Gln Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly  
20 25 30

Gly Ala Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile  
35 40 45

Trp Ala Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val  
50 55 60

Ile Thr Leu Tyr Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe  
65 70 75 80

Lys Gln Pro Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly  
85 90 95

Cys Ser Cys Arg Phe Pro Glu Glu Glu Gly Gly Cys Glu Leu  
100 105 110

<210> 590  
<211> 336  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8/CD28 sequence - N CD8 transmembrane CD28C

<400> 590  
acgacaaccc cggccccccag accaccaacg ccagccccca ccatcgccag ccaaccctg 60  
tctctgagac cagaagcctg taggcctgcc gccggtggag ctgtgcacac aagaggactg 120  
gatttcgcct gtgatatcta catttgggcc ccgctcgag gcacatgtgg agtgctcctc 180  
ctctccctgg tgattaccct gtactgcaga agcaagcggc ctcggctcct gcattctgat 240  
tacatgaaca tgacccaag aagaccaggc cccaccagga aacattacca gccctacgct 300  
ccgccacgcg acttcgctgc ctaccggtcc tgataa 336

<210> 591  
<211> 110  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CD8/CD28 sequence - N CD8 transmembrane CD28C

<400> 591

Thr Thr Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala  
1 5 10 15

Ser Gln Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly  
20 25 30

Gly Ala Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile  
35 40 45

Trp Ala Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val  
50 55 60

Ile Thr Leu Tyr Cys Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp  
65 70 75 80

Tyr Met Asn Met Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr  
85 90 95

Gln Pro Tyr Ala Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser  
100 105 110

<210> 592  
<211> 549  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8/CD3z sequence - N CD8 transmembrane CD3zetaC

<400> 592  
acgacaaccc cggccccccag accaccaacg ccagccccca ccatcgccag ccaaccctg 60  
tctctgagac cagaagcctg taggcctgcc gccggtggag ctgtgcacac aagaggactg 120  
gatttcgcct gtgatatcta catttgggcc ccgctcgag gcacatgtgg agtgctcctc 180  
ctctccctgg tgattaccct gtactgccgc gttaagttct cccgatcagc cgacgcgcct 240



gcttacaagc agggccagaa ccaactgtac aacgagctga atctcggtag acgggaagag	300
tacgacgtgt tggacaaacg gagaggccgc gaccagaaa tgggcggcaa gcctcgcagg	360
aaaaaccccc aggagggact gtacaatgag ttgcagaaag ataagatggc agaagcttat	420
agcgagatcg gaatgaaggg ggaaaggaga cgagggaaag gacacgacgg cctttatcag	480
ggcctgtcca cagcaacaaa agatacgtat gacgccctcc atatgcaggc acttcacca	540
cggtgataa	549

<210> 593  
 <211> 181  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> CD8/CD3z sequence - N CD8 transmembrane CD3zetaC  
  
 <400> 593

Thr	Thr	Thr	Pro	Ala	Pro	Arg	Pro	Pro	Thr	Pro	Ala	Pro	Thr	Ile	Ala
1				5					10					15	

Ser	Gln	Pro	Leu	Ser	Leu	Arg	Pro	Glu	Ala	Cys	Arg	Pro	Ala	Ala	Gly
			20					25					30		

Gly	Ala	Val	His	Thr	Arg	Gly	Leu	Asp	Phe	Ala	Cys	Asp	Ile	Tyr	Ile
		35					40					45			

Trp	Ala	Pro	Leu	Ala	Gly	Thr	Cys	Gly	Val	Leu	Leu	Leu	Ser	Leu	Val
	50					55					60				

Ile	Thr	Leu	Tyr	Cys	Arg	Val	Lys	Phe	Ser	Arg	Ser	Ala	Asp	Ala	Pro
65					70					75					80

Ala	Tyr	Lys	Gln	Gly	Gln	Asn	Gln	Leu	Tyr	Asn	Glu	Leu	Asn	Leu	Gly
			85						90					95	

Arg	Arg	Glu	Glu	Tyr	Asp	Val	Leu	Asp	Lys	Arg	Arg	Gly	Arg	Asp	Pro
			100					105					110		

Glu	Met	Gly	Gly	Lys	Pro	Arg	Arg	Lys	Asn	Pro	Gln	Glu	Gly	Leu	Tyr
		115						120				125			

Asn	Glu	Leu	Gln	Lys	Asp	Lys	Met	Ala	Glu	Ala	Tyr	Ser	Glu	Ile	Gly
	130					135					140				

Met	Lys	Gly	Glu	Arg	Arg	Arg	Gly	Lys	Gly	His	Asp	Gly	Leu	Tyr	Gln
145					150					155					160

Gly	Leu	Ser	Thr	Ala	Thr	Lys	Asp	Thr	Tyr	Asp	Ala	Leu	His	Met	Gln
				165					170					175	

Ala	Leu	Pro	Pro	Arg
				180

<210> 594  
<211> 672  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8/CD28/CD3z sequence - N CD8 transmembrane CD28 CD3zetaC

<400> 594  
acgacaaccc cggccccag accaccaacg ccagcccca ccatgccag ccaaccctg 60  
tctctgagac cagaagcctg taggcctgcc gccggtggag ctgtgcacac aagaggactg 120  
gatttcgcct gtgatatcta catttgggcc ccgctcgag gcacatgtgg agtgctcctc 180  
ctctccctgg tgattaccct gtactgcaga agcaagcgtt ctgggctcct gcattctgat 240  
tacaatgaac tgacccaag aagaccaggc cccaccagga aacattacca gccctacgct 300  
ccgccacgcg acttcgtgc ctaccggtcc cgcgttaagt tctccgac agccgacgcg 360  
cctgcttaca agcagggcca gaaccaactg tacaacgagc tgaatctcgg tagacgggaa 420  
gagtacgacg tgttgacaa acggagaggc cgcgaccag aaatgggcgg caagcctcgc 480  
aggaaaaacc cccaggaggg actgtacaat gagttgcaga aagataagat ggcagaagct 540  
tatagcgaga tcggaatgaa gggggaaagg agacgaggga aaggacacga cggcctttat 600  
cagggcctgt ccacagcaac aaaagatacg tatgacgcc tccatatgca ggcacttcca 660  
ccacggtgat aa 672

<210> 595  
<211> 222  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CD8/CD28/CD3z sequence - N CD8 transmembrane CD28 CD3zetaC

<400> 595

Thr Thr Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala  
1 5 10 15

Ser Gln Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly  
20 25 30

Gly Ala Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile  
35 40 45

Trp Ala Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Ser Leu Val  
50 55 60

Ile Thr Leu Tyr Cys Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp  
65 70 75 80

Tyr Met Asn Met Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr  
85 90 95

Gln Pro Tyr Ala Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val  
100 105 110

Lys Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn  
115 120 125

Gln Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val  
130 135 140

Leu Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg  
145 150 155 160

Arg Lys Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys  
165 170 175

Met Ala Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg  
180 185 190

Gly Lys Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys  
195 200 205

Asp Thr Tyr Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
210 215 220

<210> 596  
<211> 675  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> CD8/41BB/CD3z sequence - N CD8 transmembrane 41BB CD3zetaC

<400> 596  
acgacaaccc cggccccag accaccaacg ccagccccc ccatcgccag ccaacccctg 60  
tctctgagac cagaagcctg taggcctgcc gccggtggag ctgtgcacac aagaggactg 120  
gatttcgcct gtgatatcta catttgggcc ccgctcgag gcacatgtgg agtgctcctc 180  
ctctccctgg tgattaccct gtactgcaa aggggccgca aaaaactcct ttacattttt 240  
aagcagcctt ttatgaggcc agtacagacg actcaagagg aagacgggtg ctcatgccgc 300  
tttctgagg aggaggaagg aggggtgcgaa ctgcgcgtta agttctcccg atcagccgac 360  
gcgctgctt acaagcaggg ccagaaccaa ctgtacaacg agctgaatct cggtagacgg 420  
gaagagtacg acgtgttga caaacggaga ggccgcgacc cagaaatggg cggcaagcct 480  
cgcaggaaaa acccccagga gggactgtac aatgagttgc agaaagataa gatggcagaa 540  
gcttatagcg agatcggaat gaagggggaa aggagacgag ggaaaggaca cgacggcctt 600  
tatcagggcc tgtccacagc aacaaaagat acgtatgacg ccctccatat gcaggcactt 660  
ccaccacggt gataa 675

<210> 597  
<211> 223

<212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> CD8/41BB/CD3z sequence - N CD8 transmembrane 41BB CD3zetaC  
  
 <400> 597

Thr Thr Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala  
 1 5 10 15

Ser Gln Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly  
 20 25 30

Gly Ala Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile  
 35 40 45

Trp Ala Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val  
 50 55 60

Ile Thr Leu Tyr Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe  
 65 70 75 80

Lys Gln Pro Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly  
 85 90 95

Cys Ser Cys Arg Phe Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg  
 100 105 110

Val Lys Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln  
 115 120 125

Asn Gln Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp  
 130 135 140

Val Leu Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro  
 145 150 155 160

Arg Arg Lys Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp  
 165 170 175

Lys Met Ala Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg  
 180 185 190

Arg Gly Lys Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr  
 195 200 205

Lys Asp Thr Tyr Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
 210 215 220

<210> 598  
 <211> 798  
 <212> DNA  
 <213> Artificial Sequence

<220>

<223> CD8/CD28/41BB/CD3z sequence - N CD8 transmembrane CD28 41BB  
CD3zetaC

<400> 598

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acgacaaccc cggccccag accaccaacg ccagccccc ccatcgccag ccaaccctg      60
tctctgagac cagaagcctg taggcctgcc gccggtggag ctgtgcacac aagaggactg    120
gatttcgcct gtgatatcta catttgggcc ccgctcgag gcacatgtgg agtgctcctc     180
ctctccctgg tgattaccct gtactgcaga agcaagcggc ctcggctcct gcattctgat     240
tacatgaaca tgacccaag aagaccaggc cccaccagga aacattacca gccctacgct      300
ccgccacgcg acttcgtgc ctaccggtcc aaaaggggcc gcaaaaaact cttttacatt     360
ttaaagcagc cttttatgag gccagtacag acgactcaag aggaagacgg gtgctcatgc     420
cgctttcctg aggaggagga aggagggtgc gaactgcgcg ttaagttctc ccgatcagcc     480
gacgcgcctg cttacaagca gggccagaac caactgtaca acgagctgaa tctcggtaga     540
cggaagagt acgacgtgtt ggacaaacgg agaggccgcg acccagaaat gggcggcaag     600
cctcgcagga aaaaccccca ggagggactg tacaatgagt tgcagaaaga taagatggca     660
gaagcttata gcgagatcgg aatgaagggg gaaaggagac gagggaaagg acacgacggc     720
ctttatcagg gcctgtccac agcaacaaaa gatacgtatg acgccctcca tatgcaggca     780
cttcaccac ggtgataa                                     798
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<210> 599

<211> 264

<212> PRT

<213> Artificial Sequence

<220>

<223> CD8/CD28/41BB/CD3z sequence - N CD8 transmembrane CD28 41BB  
CD3zetaC

<400> 599

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Thr Thr Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala
1          5          10         15
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```
Ser Gln Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly
20        25        30
```

```
Gly Ala Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile
35        40        45
```

```
Trp Ala Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val
50        55        60
```

```
Ile Thr Leu Tyr Cys Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp
65        70        75        80
```

```
Tyr Met Asn Met Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr
85        90        95
```

Gln Pro Tyr Ala Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser Lys Arg  
 100 105 110  
 Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe Met Arg Pro  
 115 120 125  
 Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe Pro Glu  
 130 135 140  
 Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser Arg Ser Ala  
 145 150 155 160  
 Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu Leu  
 165 170 175  
 Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg Gly  
 180 185 190  
 Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln Glu  
 195 200 205  
 Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Tyr Ser  
 210 215 220  
 Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp Gly  
 225 230 235 240  
 Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr Asp Ala Leu  
 245 250 255  
 His Met Gln Ala Leu Pro Pro Arg  
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<210> 608

<211> 1482

<212> DNA

<213> Artificial Sequence

<220>

<223> CART C2 CD8/CD8/CD28/CD3z sequence - NCD81shuMNC2scFvCD8ecd  
fragment CD8 transmembrane CD28 CD3zetaC

<400> 608

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cctgaagtgc agctcgtaga gagggggg ggactggtga agcccgttg aagcctcaga	120
ctcagttgag ccgcctcagg tttcactttt tcaggttacg ccatgtcctg ggtaagacag	180
gcaccgggga aaggactcga gtgggtgtct actatcagct caggaggcac ttatatatat	240
tatcctgact ctgtaaaagg ccgatttacg atttctcgcg acaatgcaa gaactccctc	300
tacctcaaaa tgaacagtct tagggcagaa gacactgctg tatactattg tgcacgcctc	360
ggcgccgaca actactacga gtactttgac gtgtggggga aagggactac cgtgacagtt	420
tcaagcggag gaggtggctc aggtggaggc gggtcagggg ggggaggaag tgatattgtg	480
ctcacacaat cccagcctc cctggctgtg tctcccgcc aacgcgctac aattacatgt	540
cgggcctcca aaagcgtgag caccagcggc tacagctaca tgcactggta tcaacagaaa	600
ccaggacaac ccccaaact gttgatttat ctgcgttcaa acttgagtc cggcgtgcct	660
gcgcgctttt caggagtgag gagcggcaca gattttacgc tgactatcaa cccgtagaa	720
gcaaacgata cagcgaatta ttattgtcaa cattcccggg aactcccctt tacgttcggc	780
gggggcacaa aggtcgaaat taagagaacc acgacaaccc cggccccag accaccaacg	840
ccagccccc ccatcgccag ccaaccctg tctctgagac cagaagcctg taggcctgcc	900
gccggtggag ctgtgcacac aagaggactg gatttcgcct gtgatatcta catttgggcc	960
ccgctcgag gcacatgtgg agtgctcctc ctctccctgg tgattaccct gtactgcaga	1020
agcaagcggc ctggctcct gcattctgat tacatgaaca tgacccaag aagaccaggc	1080
cccaccagga aacattacca gccctacgct ccgccacgcg acttcgctgc ctaccgtcc	1140
cgcgtaagt tctccgatc agccgacgcg cctgcttaca agcagggcc gaaccaactg	1200
tacaacgagc tgaatctcgg tagacgggaa gactacgacg tggttgacaa acggagaggc	1260
cgcgaccag aaatgggcgg caagcctcgc agggaaaacc cccaggaggg actgtacaat	1320

gagttgcaga aagataagat ggcagaagct tatagcgaga tcggaatgaa gggggaaagg 1380  
agacgaggga aaggacacga cggcctttat cagggcctgt ccacagcaac aaaagatacg 1440  
tatgacgcc tccatatgca ggcacttcca ccacggtgat aa 1482

<210> 609  
<211> 492  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> CART C2 CD8/CD8/CD28/CD3z sequence - NCD81shuMNC2scFvCD8ecd  
fragment CD8 transmembrane CD28 CD3zetaC

<400> 609

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Gly Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Gly Leu Glu Trp Val Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr  
65 70 75 80

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95

Lys Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110

Ala Val Tyr Tyr Cys Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr  
115 120 125

Phe Asp Val Trp Gly Lys Gly Thr Thr Val Thr Val Ser Ser Gly Gly  
130 135 140

Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val  
145 150 155 160

Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala  
165 170 175

Thr Ile Thr Cys Arg Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser  
180 185 190

Tyr Met His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu  
195 200 205



Ile Tyr Leu Ala Ser Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser  
210 215 220

Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu  
225 230 235 240

Ala Asn Asp Thr Ala Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro  
245 250 255

Phe Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr  
260 265 270

Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln  
275 280 285

Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala  
290 295 300

Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala  
305 310 315 320

Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr  
325 330 335

Leu Tyr Cys Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met  
340 345 350

Asn Met Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro  
355 360 365

Tyr Ala Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe  
370 375 380

Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu  
385 390 395 400

Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp  
405 410 415

Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys  
420 425 430

Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala  
435 440 445

Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys  
450 455 460

Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr  
465 470 475 480

Tyr Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 610  
<211> 1485  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CART C2 CD8/CD8/41BB/CD3z sequence #13 - NCD8lshuMNC2scFvCD8ecd  
fragment CD8 transmembrane 41BB CD3zetaC

<400> 610  
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cctgaagtgc agctcgtaga gaggggcggg ggactggatg agcccggatg aagcctcaga 120  
ctcagttgca ccgctcagg ttctactttt tcaggttacg ccatgtcctg ggtaagacag 180  
gcaccgggga aaggactcga gtgggtgtct actatcagct caggaggcac ttatatatat 240  
tatcctgact ctgtaaaagg ccgatttacg atttctcgcg acaatgcaa gaactccctc 300  
tacctccaaa tgaacagtct tagggcagaa gacactgctg tatactattg tgcacgcctc 360  
ggcggcgaca actactacga gtactttgac gtgtggggga aagggactac cgtgacagtt 420  
tcaagcggag gagggtgctc aggtggaggc gggtcagggg ggggaggaag tgatattgtg 480  
ctcacacaat ccccgacctc cctggctgtg tctcccggcc aacgcgtac aattacatgt 540  
cgggcctcca aaagcgtgag caccagcggc tacagctaca tgcactggta tcaacagaaa 600  
ccaggacaac ccccaaaact gttgatttat ctgcgttcaa acttgagatc cggcgtgcct 660  
gcgcgctttt caggagtgag gagcggcaca gattttacgc tgactatcaa cccgtagaa 720  
gcaaacgata cagcgaatta ttattgtcaa cattcccggg aactcccctt tacgttcggc 780  
gggggcacaa aggtcgaaat taagagaacc acgacaacc cggccccag accaccaacg 840  
ccagccccca ccatcgccag ccaaccctg tctctgagac cagaagcctg taggcctgcc 900  
gccggtggag ctgtgcacac aagaggactg gatttcgcct gtgatatacta catttgggcc 960  
ccgctcgcag gcacatgtgg agtgctcctc ctctccctgg tgattaccct gtactgcaaa 1020  
aggggccgca aaaaactcct ttacattttt aagcagcctt ttatgaggcc agtacagacg 1080  
actcaagagg aagacgggtg ctcatgccgc ttctctgagg aggaggaagg aggggtgcgaa 1140  
ctgcgcgtta agttctcccg atcagccgac gcgcctgctt acaagcaggg ccagaaccaa 1200  
ctgtacaacg agctgaatct cggtagacgg gaagagtacg acgtgttgga caaacggaga 1260  
ggccgcgacc cagaaatggg cggcaagcct cgcaggaaaa acccccagga gggactgtac 1320  
aatgagttgc agaaagataa gatggcagaa gcttatagcg agatcggaa gaagggggaa 1380  
aggagacgag ggaaaggaca cgacggcctt tatcagggcc tgtccacagc aaaaaagat 1440  
acgtatgacg ccctccatat gcaggcactt ccaccacggt gataa 1485

<210> 611  
<211> 493  
<212> PRT  
<213> Artificial Sequence

<220>

<223> CART C2 CD8/CD8/41BB/CD3z sequence #13 - NCD81shuMNC2scFvCD8ecd  
fragment CD8 transmembrane 41BB CD3zetaC

<400> 611

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Gly Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Gly Leu Glu Trp Val Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr  
65 70 75 80

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95

Lys Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110

Ala Val Tyr Tyr Cys Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr  
115 120 125

Phe Asp Val Trp Gly Lys Gly Thr Thr Val Thr Val Ser Ser Gly Gly  
130 135 140

Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Ser Asp Ile Val  
145 150 155 160

Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala  
165 170 175

Thr Ile Thr Cys Arg Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser  
180 185 190

Tyr Met His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu  
195 200 205

Ile Tyr Leu Ala Ser Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser  
210 215 220

Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu  
225 230 235 240

Ala Asn Asp Thr Ala Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro  
245 250 255

Phe Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr  
 260 265 270

Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln  
 275 280 285

Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala  
 290 295 300

Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala  
 305 310 315 320

Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr  
 325 330 335

Leu Tyr Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln  
 340 345 350

Pro Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser  
 355 360 365

Cys Arg Phe Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys  
 370 375 380

Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln  
 385 390 395 400

Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu  
 405 410 415

Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg  
 420 425 430

Lys Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met  
 435 440 445

Ala Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly  
 450 455 460

Lys Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp  
 465 470 475 480

Thr Tyr Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
 485 490

<210> 612

<400> 612  
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<210> 613

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<210> 619

<400> 619  
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<210> 620

<211> 43

<212> PRT

<213> Artificial Sequence

<220>

<223> MUC1 truncated extra cellular domain sequence

<400> 620

Ser Asn Ile Lys Phe Arg Pro Gly Ser Val Val Val Gln Leu Thr Leu  
1 5 10 15

Ala Phe Arg Glu Gly Thr Ile Asn Val His Asp Val Glu Thr Gln Phe  
20 25 30

Asn Gln Tyr Lys Thr Glu Ala Ala Ser Arg Tyr  
35 40

<210> 621

<211> 35

<212> PRT

<213> Artificial Sequence

<220>

<223> MUC1 truncated extra cellular domain sequence

<400> 621

Ser Val Val Val Gln Leu Thr Leu Ala Phe Arg Glu Gly Thr Ile Asn  
1 5 10 15

Val His Asp Val Glu Thr Gln Phe Asn Gln Tyr Lys Thr Glu Ala Ala  
20 25 30

Ser Arg Tyr  
35

<210> 622  
<211> 24  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> MUC1 truncated extra cellular domain sequence

<400> 622

Val Gln Leu Thr Leu Ala Phe Arg Glu Gly Thr Ile Asn Val His Asp  
1 5 10 15

Val Glu Thr Gln Phe Asn Gln Tyr  
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<210> 623  
<211> 24  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> MUC1 truncated extra cellular domain sequence

<400> 623

Ser Asn Ile Lys Phe Arg Pro Gly Ser Val Val Val Gln Leu Thr Leu  
1 5 10 15

Ala Phe Arg Glu Gly Thr Ile Asn  
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<210> 624  
<211> 28  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Primer

<400> 624  
attctaagct tgggccacca tggaactg 28

<210> 625  
<211> 41  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Primer

<400> 625  
tctagagttt aaacttacta ttaccgcgga gacagggaga g 41

<210> 626

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<210>	627	
<211>	26	
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<223>	Primer	
<400>	627	
	tagaaggcac agtcgaggct gatcag	26
<210>	628	
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<400>	628	
	attctaagct tgggccacca tggaagc	27
<210>	629	
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<223>	Primer	
<400>	629	
	tctagagttt aaacttacta acactctccc ctgttgaagc	40
<210>	630	
<211>	43	
<212>	DNA	
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<400>	630	
	agtatggccc agccggccga aattgtgttg acacagtctc cag	43
<210>	631	
<211>	26	
<212>	DNA	
<213>	Artificial Sequence	
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<223>	Primer	
<400>	631	
	tagaaggcac agtcgaggct gatcag	26

<210> 632  
<211> 33  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Primer  
  
<400> 632  
actgtcatat ggaggtgcag ctggtggagt ctg 33

<210> 633  
<211> 37  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Primer  
  
<400> 633  
actgtctcga gtttaatttc cactttggtg ccgctgc 37

<210> 634  
<211> 33  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Primer  
  
<400> 634  
actgtcatat ggaggtgcag ctggtggagt ctg 33

<210> 635  
<211> 37  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Primer  
  
<400> 635  
actgtaccgg ttttaatttc cactttggtg ccgctgc 37

<210> 636  
<211> 29  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Primer  
  
<400> 636  
cttcttctc aggagcaagc tcaccgtgg 29

<210> 637  
<211> 18  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Primer



<400> 637	
gagccgtcgg agtccagc	18
<210> 638	
<211> 18	
<212> DNA	
<213> Artificial Sequence	
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<223> Primer	
<400> 638	
gcacctgaac tcctgggg	18
<210> 639	
<211> 22	
<212> DNA	
<213> Artificial Sequence	
<220>	
<223> Primer	
<400> 639	
tttaatttcc actttggtgc cg	22
<210> 640	
<211> 32	
<212> DNA	
<213> Artificial Sequence	
<220>	
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<400> 640	
cgcggtctagc ttaagcttgg taccgagggc ca	32
<210> 641	
<211> 36	
<212> DNA	
<213> Artificial Sequence	
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cgcggtcgcc gcctgatcag cgggtttaaa cttatc	36
<210> 642	
<211> 2151	
<212> DNA	
<213> Artificial Sequence	
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cccagacagc gccagtccac ctttgtgctc ttccctggag acctgagaac caatctcacc	120
gacaggcagc tggcagagga atacctgtac cgctatggtt acactcgggt ggcagagatg	180
cgtggagagt cgaaatctct ggggcctgcg ctgctgcttc tccagaagca actgtccctg	240

cccgagaccg gtgagctgga tagcgccacg ctgaaggcca tgcgaacccc acggtgcggg	300
gtcccagacc tgggcagatt ccaaaccttt gagggcgacc tcaagtggca ccaccacaac	360
atcacctatt ggatccaaaa ctactcgga gacttgccgc gggcggatgat tgacgacgcc	420
tttggccgcg ccttcgact gtggagcgcg gtgacgccgc tcaccttcac tcgcgtgtac	480
agccgggacg cagacatcgt catccagttt ggtgtcgcgg agcacggaga cgggtatccc	540
ttcgacggga aggacgggct cctggcacac gcctttcctc ctggccccgg cattcaggga	600
gacgcccatt tcgacgatga cgagttgtgg tccctgggca agggcgctgt ggttccaact	660
cggtttgaa acgcagatgg cgcggcctgc cactttccct tcatcttcga gggccgctcc	720
tactctgcct gcaccaccga cggtcgctcc gacggcttgc cctggtgcag taccacggcc	780
aactacgaca ccgacgacc gtttggcttc tgccccagcg agagactcta caccaggac	840
ggcaatgctg atgggaaacc ctgccagttt ccattcatct tccaaggcca atcctactcc	900
gcctgcacca cggacggtcg ctccgacggc taccgctggt gcgccaccac cgccaactac	960
gaccgggaca agctcttcgg cttctgcccc acccgagctg actcgacggt gatggggggc	1020
aactcggcgg gggagctgtg cgtcttcccc ttcactttcc tgggtaagga gtactcgacc	1080
tgtaccagcg agggccgcgg agatgggcgc ctctggtgcg ctaccacctc gaactttgac	1140
agcgacaaga agtggggctt ctgcccggac caaggataca gtttgttcct cgtggcggcg	1200
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cctatgtacc gttcactga ggggcccc ttgcataagg acgacgtgaa tggcatccgg	1320
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<210> 643  
 <211> 715  
 <212> PRT

<213> Artificial Sequence

<220>

<223> MMP9 sequence

<400> 643

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Cys Phe Ala Ala Pro Arg Gln Arg Gln Ser Thr Leu Val Leu Phe Pro  
20 25 30

Gly Asp Leu Arg Thr Asn Leu Thr Asp Arg Gln Leu Ala Glu Glu Tyr  
35 40 45

Leu Tyr Arg Tyr Gly Tyr Thr Arg Val Ala Glu Met Arg Gly Glu Ser  
50 55 60

Lys Ser Leu Gly Pro Ala Leu Leu Leu Leu Gln Lys Gln Leu Ser Leu  
65 70 75 80

Pro Glu Thr Gly Glu Leu Asp Ser Ala Thr Leu Lys Ala Met Arg Thr  
85 90 95

Pro Arg Cys Gly Val Pro Asp Leu Gly Arg Phe Gln Thr Phe Glu Gly  
100 105 110

Asp Leu Lys Trp His His His Asn Ile Thr Tyr Trp Ile Gln Asn Tyr  
115 120 125

Ser Glu Asp Leu Pro Arg Ala Val Ile Asp Asp Ala Phe Ala Arg Ala  
130 135 140

Phe Ala Leu Trp Ser Ala Val Thr Pro Leu Thr Phe Thr Arg Val Tyr  
145 150 155 160

Ser Arg Asp Ala Asp Ile Val Ile Gln Phe Gly Val Ala Glu His Gly  
165 170 175

Asp Gly Tyr Pro Phe Asp Gly Lys Asp Gly Leu Leu Ala His Ala Phe  
180 185 190

Pro Pro Gly Pro Gly Ile Gln Gly Asp Ala His Phe Asp Asp Asp Glu  
195 200 205

Leu Trp Ser Leu Gly Lys Gly Val Val Val Pro Thr Arg Phe Gly Asn  
210 215 220

Ala Asp Gly Ala Ala Cys His Phe Pro Phe Ile Phe Glu Gly Arg Ser  
225 230 235 240

Tyr Ser Ala Cys Thr Thr Asp Gly Arg Ser Asp Gly Leu Pro Trp Cys  
245 250 255

Ser Thr Thr Ala Asn Tyr Asp Thr Asp Asp Arg Phe Gly Phe Cys Pro  
 260 265 270

Ser Glu Arg Leu Tyr Thr Gln Asp Gly Asn Ala Asp Gly Lys Pro Cys  
 275 280 285

Gln Phe Pro Phe Ile Phe Gln Gly Gln Ser Tyr Ser Ala Cys Thr Thr  
 290 295 300

Asp Gly Arg Ser Asp Gly Tyr Arg Trp Cys Ala Thr Thr Ala Asn Tyr  
 305 310 315 320

Asp Arg Asp Lys Leu Phe Gly Phe Cys Pro Thr Arg Ala Asp Ser Thr  
 325 330 335

Val Met Gly Gly Asn Ser Ala Gly Glu Leu Cys Val Phe Pro Phe Thr  
 340 345 350

Phe Leu Gly Lys Glu Tyr Ser Thr Cys Thr Ser Glu Gly Arg Gly Asp  
 355 360 365

Gly Arg Leu Trp Cys Ala Thr Thr Ser Asn Phe Asp Ser Asp Lys Lys  
 370 375 380

Trp Gly Phe Cys Pro Asp Gln Gly Tyr Ser Leu Phe Leu Val Ala Ala  
 385 390 395 400

His Glu Phe Gly His Ala Leu Gly Leu Asp His Ser Ser Val Pro Glu  
 405 410 415

Ala Leu Met Tyr Pro Met Tyr Arg Phe Thr Glu Gly Pro Pro Leu His  
 420 425 430

Lys Asp Asp Val Asn Gly Ile Arg His Leu Tyr Gly Pro Arg Pro Glu  
 435 440 445

Pro Glu Pro Arg Pro Pro Thr Thr Thr Thr Pro Gln Pro Thr Ala Pro  
 450 455 460

Pro Thr Val Cys Pro Thr Gly Pro Pro Thr Val His Pro Ser Glu Arg  
 465 470 475 480

Pro Thr Ala Gly Pro Thr Gly Pro Pro Ser Ala Gly Pro Thr Gly Pro  
 485 490 495

Pro Thr Ala Gly Pro Ser Thr Ala Thr Thr Val Pro Leu Ser Pro Val  
 500 505 510

Asp Asp Ala Cys Asn Val Asn Ile Phe Asp Ala Ile Ala Glu Ile Gly  
 515 520 525

Asn Gln Leu Tyr Leu Phe Lys Asp Gly Lys Tyr Trp Arg Phe Ser Glu  
530 535 540

Gly Arg Gly Ser Arg Pro Gln Gly Pro Phe Leu Ile Ala Asp Lys Trp  
545 550 555 560

Pro Ala Leu Pro Arg Lys Leu Asp Ser Val Phe Glu Glu Arg Leu Ser  
565 570 575

Lys Lys Leu Phe Phe Phe Ser Gly Arg Gln Val Trp Val Tyr Thr Gly  
580 585 590

Ala Ser Val Leu Gly Pro Arg Arg Leu Asp Lys Leu Gly Leu Gly Ala  
595 600 605

Asp Val Ala Gln Val Thr Gly Ala Leu Arg Ser Gly Arg Gly Lys Met  
610 615 620

Leu Leu Phe Ser Gly Arg Arg Leu Trp Arg Phe Asp Val Lys Ala Gln  
625 630 635 640

Met Val Asp Pro Arg Ser Ala Ser Glu Val Asp Arg Met Phe Pro Gly  
645 650 655

Val Pro Leu Asp Thr His Asp Val Phe Gln Tyr Arg Glu Lys Ala Tyr  
660 665 670

Phe Cys Gln Asp Arg Phe Tyr Trp Arg Val Ser Ser Arg Ser Glu Leu  
675 680 685

Asn Gln Val Asp Gln Val Gly Tyr Val Thr Tyr Asp Ile Leu Gln Cys  
690 695 700

Pro Glu Asp Asp Tyr Lys Asp Asp Asp Asp Lys  
705 710 715

<210> 644

<211> 1062

<212> DNA

<213> Artificial Sequence

<220>

<223> MMP9 catalytic domain

<400> 644

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gcactgtgga gcgcggtgac gccgctcacc ttcactcgcg tgtacagccg ggacgcagac 180

atcgatcatcc agtttggtgt cgcgagacac ggagacgggt atcccttcga cggaaggac 240

gggctcctgg cacacgcctt tcctcctggc cccggcattc agggagacgc ccatttcgac 300

gatgacgagt tgtggtccct gggcaagggc gtcgtggttc caactcggtt tggaaacgca 360

gatggcgcgg cctgccactt ccccttcata ttcgagggcc gctcctactc tgcctgcacc 420  
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<210> 645  
<211> 352  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> MMP9 catalytic domain

<400> 645

Met Phe Gln Thr Phe Glu Gly Asp Leu Lys Trp His His His Asn Ile  
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Thr Tyr Trp Ile Gln Asn Tyr Ser Glu Asp Leu Pro Arg Ala Val Ile  
20 25 30

Asp Asp Ala Phe Ala Arg Ala Phe Ala Leu Trp Ser Ala Val Thr Pro  
35 40 45

Leu Thr Phe Thr Arg Val Tyr Ser Arg Asp Ala Asp Ile Val Ile Gln  
50 55 60

Phe Gly Val Ala Glu His Gly Asp Gly Tyr Pro Phe Asp Gly Lys Asp  
65 70 75 80

Gly Leu Leu Ala His Ala Phe Pro Pro Gly Pro Gly Ile Gln Gly Asp  
85 90 95

Ala His Phe Asp Asp Asp Glu Leu Trp Ser Leu Gly Lys Gly Val Val  
100 105 110

Val Pro Thr Arg Phe Gly Asn Ala Asp Gly Ala Ala Cys His Phe Pro  
115 120 125

Phe Ile Phe Glu Gly Arg Ser Tyr Ser Ala Cys Thr Thr Asp Gly Arg

130	135	140
Ser Asp Gly Leu Pro Trp Cys Ser Thr Thr Ala Asn Tyr Asp Thr Asp 145 150 155 160		
Asp Arg Phe Gly Phe Cys Pro Ser Glu Arg Leu Tyr Thr Gln Asp Gly 165 170 175		
Asn Ala Asp Gly Lys Pro Cys Gln Phe Pro Phe Ile Phe Gln Gly Gln 180 185 190		
Ser Tyr Ser Ala Cys Thr Thr Asp Gly Arg Ser Asp Gly Tyr Arg Trp 195 200 205		
Cys Ala Thr Thr Ala Asn Tyr Asp Arg Asp Lys Leu Phe Gly Phe Cys 210 215 220		
Pro Thr Arg Ala Asp Ser Thr Val Met Gly Gly Asn Ser Ala Gly Glu 225 230 235 240		
Leu Cys Val Phe Pro Phe Thr Phe Leu Gly Lys Glu Tyr Ser Thr Cys 245 250 255		
Thr Ser Glu Gly Arg Gly Asp Gly Arg Leu Trp Cys Ala Thr Thr Ser 260 265 270		
Asn Phe Asp Ser Asp Lys Lys Trp Gly Phe Cys Pro Asp Gln Gly Tyr 275 280 285		
Ser Leu Phe Leu Val Ala Ala His Glu Phe Gly His Ala Leu Gly Leu 290 295 300		
Asp His Ser Ser Val Pro Glu Ala Leu Met Tyr Pro Met Tyr Arg Phe 305 310 315 320		
Thr Glu Gly Pro Pro Leu His Lys Asp Asp Val Asn Gly Ile Arg His 325 330 335		
Leu Tyr Gly Pro Arg Pro Glu Pro Asp Tyr Lys Asp Asp Asp Asp Lys 340 345 350		

<210> 646  
 <211> 2186  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> NFATc1 Promoter (NFATc1P)

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cagagggaaa cagacccggg attccgtcac ccgggcgggg ggataaggac ggctttgaga	240
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gaggcggcag cggaggaagg ggaggggagg gaggaggagg aagagccggc cgccccgcc	360
ccggccccgg ctctcagga gccaaaggga gcctcgccag gtcgggtccc ggctcagga	420
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<210> 647  
<211> 4349  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> NFATc1PMMP9

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 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> NFATc1PMMP9cat

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ccaaaccttt gagggcgacc tcaagtggca ccaccacaac atcacctatt ggatccaaaa	240
ctactcggaa gacttgccgc gggcgggtgat tgacgacgcc tttgcccgcg ctttcgcact	300
gtggagcgcg gtgacgccgc tcaccttcac tcgcgtgtac agccgggacg cagacatcgt	360
catccagttt ggtgtcgcgg agcacggaga cgggtatccc ttcgacggga aggacgggct	420
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cgagttgtgg tccctgggca agggcgctcgt ggttccaact cggtttgaa acgcagatgg	540
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cggtcgctcc gacggcttgc cctggtgcag taccacggcc aactacgaca ccgacgaccg	660
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 <223> C2 scFv

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cccgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat	240
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Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly



195

200

205

Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu Ala Asn Asp Thr Ala  
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Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro Phe Thr Phe Gly Gly  
 225 230 235 240

Gly Thr Lys Val Glu Ile Lys Arg Thr  
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&lt;210&gt; 656

&lt;211&gt; 72

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; CD8 transmembrane domain

&lt;400&gt; 656

atctacatct gggcgccctt ggccgggact tgtgggggtcc ttctcctgtc actggttatc 60

accctttact gc 72

&lt;210&gt; 657

&lt;211&gt; 24

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; CD8 transmembrane domain

&lt;400&gt; 657

Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu  
 1 5 10 15

Ser Leu Val Ile Thr Leu Tyr Cys  
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&lt;210&gt; 658

&lt;211&gt; 126

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; 41BB domain

&lt;400&gt; 658

aaacggggca gaaagaaact cctgtatata ttcaacaac catttatgag accagtacaa 60

actactcaag aggaagatgg ctgtagctgc cgatttcag aagaagaaga aggaggatgt 120

gaactg 126

&lt;210&gt; 659

&lt;211&gt; 42

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

<223> 41BB domain

<400> 659

Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe Met  
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Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe  
20 25 30

Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu  
35 40

<210> 660

<211> 336

<212> DNA

<213> Artificial Sequence

<220>

<223> CD3zeta domain

<400> 660

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cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgtacaat 180  
gaactgcaga aagataagat ggcggaggcc tacagtgaga ttgggatgaa aggcgagcgc 240  
cggagggggca aggggcacga tggcctttac cagggtctca gtacagccac caaggacacc 300  
tacgacgcc ttcacatgca ggccctgccc cctcgc 336

<210> 661

<211> 112

<212> PRT

<213> Artificial Sequence

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<223> CD3zeta domain

<400> 661

Arg Val Lys Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly  
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Gln Asn Gln Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr  
20 25 30

Asp Val Leu Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys  
35 40 45

Pro Arg Arg Lys Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys  
50 55 60

Asp Lys Met Ala Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg  
65 70 75 80

Arg Arg Gly Lys Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala

Thr	Lys	Asp	Thr	Tyr	Asp	Ala	Leu	His	Met	Gln	Ala	Leu	Pro	Pro	Arg
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<210> 718  
<211> 1485  
<212> DNA  
<213> Artificial Sequence

<220>

<223> CART C2 CD8/CD8/41BB/CD3z #44 - NCD8lshuMNC2scFvCD8ecd fragment  
CD8 transmembrane 41BB CD3zetaC

<400> 718

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gctccaggga aggggctgga gtgggtctca accattagta gtggcggaac ctacatatac	240
taccccgact cagtgaaggg ccgattcacc atctccagag acaacgcaa gaactcactg	300
tatctgcaaa tgaacagcct gagagccgag gacacggccg tgtattactg tgcgagactt	360
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1485

<210> 719

<211> 493

<212> PRT

<213> Artificial Sequence

<220>

<223> CART C2 CD8/CD8/41BB/CD3z #44 - NCD8lshuMNC2scFvCD8ecd fragment  
CD8 transmembrane 41BB CD3zetaC

<400> 719

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
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His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Gly Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Gly Leu Glu Trp Val Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr  
65 70 75 80

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95

Lys Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110

Ala Val Tyr Tyr Cys Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr  
115 120 125

Phe Asp Val Trp Gly Lys Gly Thr Thr Val Thr Val Ser Ser Gly Gly  
130 135 140

Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val  
145 150 155 160

Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala  
165 170 175

Thr Ile Thr Cys Arg Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser  
180 185 190

Tyr Met His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu  
195 200 205

Ile Tyr Leu Ala Ser Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser  
210 215 220



Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu  
225 230 235 240

Ala Asn Asp Thr Ala Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro  
245 250 255

Phe Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr  
260 265 270

Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln  
275 280 285

Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala  
290 295 300

Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala  
305 310 315 320

Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr  
325 330 335

Leu Tyr Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln  
340 345 350

Pro Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser  
355 360 365

Cys Arg Phe Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys  
370 375 380

Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln  
385 390 395 400

Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu  
405 410 415

Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg  
420 425 430

Lys Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met  
435 440 445

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gcagacagga aaaggagct tttctgcatg gggtgaaaaa attatttatt gaaggaggag	300
gaggcggcag cggaggaagg ggagggggcg gaggaggagg aagagccggc cgccccgcc	360
ccggccccgg ctctcagga gccaaggga gcctcgccag gtcggtccc ggctcgagga	420
ccgcggctgg ggtcgagggg ctcagtctcc cacgtgaccg gctgggcgcg ccccgccaga	480
ccggcctcg ggattccctc ctcccggcga gtctccgcc gcccgtcct ggaggtgggg	540
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ggggacactc gcggcgggaa gatttgagg ggaggggagg gggagggcg tggggcgcg	660
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cggacgggct ctggccgcgc accttcgagg gctctgcag cccgaccgc ctccccggc	1020
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cctcctgggc cccgcagcag acacgagttt agcctttggg tttagtttaa atcacataag	1140
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catccaagcc aacaagaaaa tgcggtgccg tgttggcagc tgagctgcgc ccgaagagac	1260
gcaggagac gtaagagagg aaagtgtgag tggccgggg gcctcccc gtcagaagtc	1320
gcgcagtcgc gccataaaa cgccccctc gggcggctag ggcaggtgag cgcgtccccg	1380

ggcctcccca cgccggcccc tgccacagag ccgtctaggt cgagcagata tttacagaat	1440
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atgggaattt cttttctagg gcctgccagt gaagcgcttt tccaaatttc cacagcgggg	1560
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cgccccctcc cgtgcgcccc cctccccgtg cgccccccct ccccgctgcgc cccctcccc	1860
gtgcgcccc cctccccggg cgccccctc cccgggcgcc cccctcccc gtgcgcccc	1920
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cgagcggctg ccgcggcgcg gggaggggcg ggcgctcggc gactcgtccc cggggccccg	2040
cgcgggcccc ggagcaggg gcgtgatgtc acggcagggg gggggcgcg gagccgccgg	2100
gccggcgggg aggcggggga ggtgttttcc agctttaaaa aggcaggagg cagagcgcg	2160
ccctgcgtca gagcgagact cagaggctcc gaactcgccg gcggagtcgc cgcgccagat	2220
cccagcagca gggcgcg	2238

<210> 782  
 <211> 2186  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> NFATc1 Promoter fragment (P2)

<400> 782	
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cgctccgga actccctgcg cctggcgcg gccaccgtg gtcccgcaa cggcattaaa	180
cagagggaag cagaccggg attccgtcac ccggcgggg ggataaggac ggctttgaga	240
gcagacagga aaaggagct tttctgcatg gggtgaaaa attatttatt gaaggaggag	300
gaggcgagc cggaggaagg ggagggcggg gaggaggagg aagagccggc cggccccgc	360
ccggccccgg ctctcagga gccaaaggga gcctcgccag gtcggtccc ggctcagga	420
ccggcgctgg ggtcgagggg ctcagttcc cacgtgacc gctggcgcg ccccgccaga	480
ccggcctcg ggattccctc ctccggcga gtctccgcc gcccgtcct ggaggtgggg	540
agaaggaggg cggggcgggg gggacggaat ctctcccgcc caaatcctgg cccaggcct	600
ggggacactc gcggcgggaa gatttgagg ggaggggagg gggagggcg tggggcgcg	660
gcctcgtgg agtccccctg accccccgac ccccgccac cggcctgggc gtcctccgc	720
ggccccctc cccctccgg cgcccggtgc tctggggcg gtgccacgc tggctcggcg	780
ccgtaggggc ccccgaggt agagaccct ggaaatggcc tcgacgccg aggagcagg	840
cggccaccac cccgctaac cgggcacgtc tctccaggcc gaggcctgc gtggaaaagc	900

cggggttcca tttgtgctga gtcggggcgg ccgaatggag ccaggcctcg ggacgcggga	960
cggacgggct ctggccgcgc accttcgcgg gctctgcagc gcccgaccgc ctccccggc	1020
agggaggagg cgcttggtgg gggcacccac ggggcacagt gatccctggg ggtctgcgga	1080
cctcctgggc ccgcagcag acacgagttt agcctttggg tttagtttaa atcacataag	1140
ggtgtcgtgc aatcgattta tggtttctac acaccagaca ctttaacctc caaccccccc	1200
catccaagcc aacaagaaaa tgcggtgccg tggtggcagc tgagctgcgc ccgaagagac	1260
gcaggagagc gtaagagagg aaagtgtgag tggccggggg gcctcccccc gtcagaagtc	1320
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atgggaattt cttttctagg gcctgccagt gaagcgcttt tccaaatttc cacagcgggg	1560
gaagcctgcg attttacata atgacttcag catgccgggc tttctcgaca cccctccccg	1620
gccccgggc cccgcccccc gcccttttc cagcagggcc gggctccctc cggacaccgc	1680
cgtggactca ggcgtcccg ctggcccgtt cgccccgtt tccccgcca gcccagcgc	1740
ccccctgccc ggcccccgga ttccccgtt ccgcccctac gccccatcc cctccccgtg	1800
cgccccctcc cgtgcgcccc cctccccgtg cgccccctt ccccgctgcgc cccccctcc	1860
gtgcgcccc cctccccggg cgccccctc ccggggcgcc cccccctccc gtgcgcccc	1920
ccctccccgt gcgcccccc tccccgtgc cgccccgcct cttgcgcccc tgccccagg	1980
cgagcggctg ccgcggcgcg gggaggggcg ggcgtcggc gactcgtccc cggggccccg	2040
cgcgggcccc ggcagcagg gcgtgatgtc acggcaggga gggggcgcg gagccgccgg	2100
gccggcgggg aggcggggga ggtgttttcc agctttaaaa aggcaggagg cagagcgcg	2160
ccctgcgtca gagcgagact cagagg	2186

<210> 783

<211> 1645

<212> DNA

<213> Artificial Sequence

<220>

<223> NFATc1 Promoter fragment (P3)

<400> 783

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gggcgcggcc tcgtggagt cccctgacc ccccgacccc cgcccaccgg cctgggcgtc	120
ctcccgcggc cctcctccc ctccggcgcg ccggtgctct ggggcgcgtg ccacgcctgg	180
ctcggcgccg taggggcccc cgaggtaga gaccctgga aatggcctcg acgccgagg	240
agcaggcgcg ccaccacccc gctaattcgg gcacgtctct ccaggccgag gcctgcggtg	300
gaaaagccgg ggttccattt gtgctgagtc ggggcggccg aatggagcca ggcctcggga	360
cgcgggacgg acgggctctg gccgcgcacc ttgcgggct ctgcagcgcc cgaccgcctc	420

ccccggcagg gaggaggcgc ttgtgggggg caccacggg gcacagtgat ccctgggggt	480
ctgcggacct cctgggcccc gcagcagaca cgagtttagc ctttgggttt agtttaaadc	540
acataagggt gtcgtgcaat cgatttatgg tttctacaca ccagacactt taacctcaa	600
cccccccat ccaaagccaa caagaaaatg cggtgccgtg ttggcagctg agctgcgccc	660
gaagagacgc agggagacgt aagagaggaa agtgtgagtg gccggggggc ctcccccg	720
cagaagtgcg gcagtcgcgc ccataaacg cccctccgg gcggctaggg caggtgagcg	780
cgtccccggg cctccccacg ccggcccctg ccacaggccg tctaggtcga gcagatattt	840
acagaataaa aatgacaata actcgacgtc ccgggacggc cagcaatct gttagtaatt	900
tagcgggatg ggaatttcct ttctagggcc tgccagtga ggccttttcc aaatttcac	960
agcgggggaa gcctgcgatt ttacataatg acttcagcat gccgggcttt ctcgacacc	1020
ctccccggc cccggcccc gcccccgc cttttccag cagggccggg ctccctccg	1080
acaccgcgt ggactcaggc gtccgtctg gccgttcgc cccgtttcc ccgcccagc	1140
ccagcgcgcc cctgcccggc ccccgattc cccgttccc cccctacgc cccatcccct	1200
ccccgtgcgc ccctccccgt gcgccccct cccgtgcgc cccccctcc cgtgcgccc	1260
cctccccgtg cgccccct cccggggcg cccctccc gggcgcccc cctccccgtg	1320
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ccccaggcga gcggctgccg cggcgcgggg agggcgggc gctcggcgac tcgtccccg	1440
ggccccgcgc gggcccggg agcagggcg tgatgtcac gcaggaggg ggcgcgggag	1500
ccgcccggcc ggcggggagg gggggagggt gttttccag tttaaaagg caggaggcag	1560
agcgcgggcc tgcgtcagag cgagactcag aggctccga ctcgccggcg gagtgcgcg	1620
gccagatccc agcagcaggg cgcgg	1645

<210> 784

<211> 927

<212> DNA

<213> Artificial Sequence

<220>

<223> pNFATMMP9cat1 gBLOCK sequence

<400> 784

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tatataagca gagctcgttt agtgaaccgt cagatcgct ggagacgcca tccacgtgt	120
tttgacctcc atagaagaca ccgggaccga tccagcctct cgacattcgt ttctagagcc	180
accatgagcc tctggcagcc cctggtcctg gtgctcctgg tgctgggctg ctgctttgct	240
ttccaaacct ttgaggcgga cctcaagtgg caccaccaca acatcaccta ttgatccaa	300
aactactcgg aagacttgcc gcgggcgggtg attgacgac cttttgccg cgccttcgca	360
ctgtggagcg cggtgacgcc gtcaccttc actcgctgt acagccggga cgcagacatc	420
gtcatccagt ttggtgtcgc ggagcacgga gacgggtatc ctttcgacgg gaaggacggg	480
ctcctggcac acgcctttcc tcctggcccc ggcatcagg gagacgcca tttcgacgat	540

gacgagttgt ggtccctggg caagggcgtc gtggttcaa ctcggtttgg aaacgcagat	600
ggcgcggcct gccacttccc cttcatcttc gagggccgct cctactctgc ctgcaccacc	660
gacggtcgct ccgacggctt gccctggtgc agtaccacgg ccaactacga caccgacgac	720
cggtttggct tctgccccag cgagagactc tacacccagg acggcaatgc tgatgggaaa	780
ccctgccagt ttccattcat cttccaaggc caatcctact ccgcctgcac cacggacggt	840
cgctccgacg gctaccgctg gtgcgccacc accgccaact acgaccggga caagctcttc	900
ggcttctgcc cgacccgagc tgactcg	927

<210> 785

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<210> 800

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<210> 801

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<210> 802

<400> 802  
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<210> 803

<400> 803  
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<210> 804

<211> 13

<212> PRT

<213> Artificial Sequence

<220>

<223> NFAT consensus sequence

<400> 804

Ala Thr Gly Gly Ala Ala Ala Ala Asn Ala Thr Cys Asn  
1 5 10

<210> 805

<211> 32

<212> DNA

<213> Artificial Sequence

<220>

<223> Current NFAT RE (Form System Biosciences. The sequence is from  
the mouse IL2 promoter.)

<400> 805

aagaggaaaa ttgtttcat acagaaggcg tt

<210> 806  
<211> 812  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse IL2 Promoter (highlighted in green the NFAT RE used,  
highlighted in yellow is the start codon)

<400> 806  
aactagagac atataaaata acaccaacat ccttagatac aacccttcct gagaatttat 60  
tggacatcat actcttttta aaaagcataa taaacatcaa gacacttaca caaaatatgt 120  
taaattaaat ttaaaacaac aacgacaaaa tagtacctca agctcaacaa gcatttttagg 180  
tgtccttagc ttactatttc tctggctaac tgtatgaagc catctatcac cctgtgtgca 240  
attagctcat tgtgtagata agaaggtaaa accatcttga aacaggaaac caatatcctt 300  
cctgtctaata caacaatct aaaagattta ttcttttcat ctatctcctc ttgcgtttgt 360  
ccaccacaac aggctgctta cagggttcagg atggttttga caaagagaac attttcatga 420  
gttacttttg tgtctccacc ccaaagagga aaatttgttt catacagaag gcgttcattg 480  
tatgaattaa aactgccacc taagtgtggg ctaacccgac caagagggat ttcacctaaa 540  
tccattcagt cagtgtatgg gggtttaaaag aaattccaga gagtcatcag aagaggaaaa 600  
acaaaaggta atgctttctg ccacacaggt agactctttg aaaatatgtg taatatgtaa 660  
aacatcgtga ccccccata ttatttttcc agcattaaca gtataaattg cctcccatgc 720  
tgaagagctg cctatcacc ttgctaata ctcctcacag tgacctcaag tcctgcaggc 780  
atgtacagca tgcagctcgc atcctgtgtc ac 812

<210> 807  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> NFAT RE (Form PRomega. The sequence is from the humane IL2  
promoter.)

<400> 807  
ggaggaaaaa ctgtttcata cagaaggcgt 30

<210> 808  
<211> 33  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Possible NFAT RE from ET1 promoter

<400> 808  
tccagggaat atcggagtag aacaagaggg atg 33

<210> 809  
<211> 34  
<212> DNA  
<213> Artificial Sequence

<220>  
 <223> Possible NFAT RE from ET1 promoter  
  
 <400> 809  
 actgttgga aacgtaaaca cgttattaaa cggt 34

<210> 810  
 <211> 20  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Possible NFAT RE from human CD3?  
  
 <400> 810  
 tccttaacgg aaaaacaaaa 20

<210> 811  
 <211> 20  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Possible NFAT RE from human CD3?  
  
 <400> 811  
 aaaggaaaa gtatatgttc 20

<210> 812  
 <211> 17  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Possible NFAT RE from human IL3 promoter  
  
 <400> 812  
 atgccatgga aagggtg 17

<210> 813  
 <211> 22  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Possible NFAT RE from human GPC6  
  
 <400> 813  
 aaggggaaat gttgagtcta ga 22

<210> 814  
 <211> 17  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Possible NFAT RE from human growth hormonereleasing hormone  
  
 <400> 814  
 aacttgaaa agcatag 17

<210> 815  
 <211> 5320

<212> DNA

<213> Artificial Sequence

<220>

<223> NFATc1 promoter large

<400> 815

ttatgccgtc tagaggagac atacttttcta ctcaaagcta cacacataga ctacaacgat	60
gggaaaagac gacacaccaa cagcgacttc aggaaagctg gagtggctgc taatgttaga	120
caaaaataggc tttttaaaaa aggtttttatt aaagaggaat gtttcgtaat gataaaagca	180
ctaattctgtg agaaagatac aacaatgata aacatacgtg cagctaataa gagagctcca	240
aaatctatga agcaaaaact cacagaatga ggggagaagc agttctacaa cagagaatgg	300
ggacttcgat actccacttt caataatgga tacaacaacc aggagataa caaggcaaca	360
gaaggcctga acaacagtat aaaccaatta gacctaccag atatctatag ctagcacact	420
ccaccaacg acagcagaat acacatttctt ctcaagcgca caagtaacat cctccaggat	480
gggcatgtt ctaggccatc aaacaaactc aggtggtttg aggccagagg cctctctttt	540
aaccaccaca ctagggcctt cggaggaggc aagcagagag ttgtcaaaga ggccctcagg	600
actgggtgca gtggctcatg actgtaatcc cagcacttta gaaggctgag gcacaaggat	660
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tgtcagcacc ttctgacttt ctttctaaag tctggggtgt tcctgaggat ccgtaagttt	1260
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gagttgcccc ggagttaag accagcctga gcaacatagc aagacctcat ctcaacaaca	1380
acaaaaatta gaaataaatt agccagggtgt ggtgacatgt gcctgtagtc ccagctactc	1440
agaaggctga ggcatgagga tcacttgggc ccaggaagtt ggggctgcag ggagccctgt	1500
tcatgccgct gcactccagc ctgcaagaca gagcagaaaa aaagaatcag gatcctgggc	1560
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ctgatccatt ggtgatctga gtttttagac atgggtggtag tccatgaggt gggtgttcat	1860

gctaagagtt tagacagggga aacctatgaa gcccttagca accctccagg gaaggggcggt	1920
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agaacacagt cgtattaagt gattttcccg agcaggaagt ggcatctggc ctgcggttca	2460
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cagaggccgg gccggggcg agaggccggg cgagctggcc gcgctctggg ccgccgcctc	3120
cggaaactcc tgcgcctggc gcgcggccac cgtggtcccc gcaacggcat taaacagagg	3180
gaaacagacc cgggattccg tcacccgggc ggggggataa ggacggcttt gagagcagac	3240
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gcagcggagg aaggggaggg gcgggaggag gaggaagagc cggccgcccc cgccccggcc	3360
ccggctcctc aggagccaag ggcagcctcg ccaggtcggg cccgggctcg aggaccgcgg	3420
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ctcgggattc cctcctcccg gcgagtctcc gcccgccccg tcctggaggt ggggagaagg	3540
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actcgcggcg ggaagatttg gaggggaggg gagggggagg ggcgtggggg cgcggcctcg	3660
ctggagtccc cctgaccccc cgacccccgc ccaccggcct gggcgtcctc ccgcggcccc	3720
tcctccccct ccggcgcccc gtgctctggg gcgcgtgcc a gcctggctc ggcgccgtag	3780
gggccccgc aggtagagac ccctggaaat ggcctcgacg ccgcaggagc gaggcggcca	3840
ccaccccgct aatccgggca cgtctctcca ggccgaggcc tgcggtggaa aagccgggggt	3900
tccatttgtg ctgagtcggg gcggccgaat ggagccaggc ctcgggacgc gggacggacg	3960

ggctctggcc gcgcaccttc gcgggctctg cagcgcccga ccgcctcccc cggcagggag	4020
gaggcgcttg tggggggcac ccacggggca cagtgatccc tgggggtctg cggacctcct	4080
gggccccga gcagacacga gtttagcctt tgggtttagt ttaaatacaca taagggtgtc	4140
gtgcaatcga tttatggttt ctacacacca gacctttta cctccaaccc ccccatcca	4200
aagccaacaa gaaaatgcgg tgccgtgttg gcagctgagc tgcgcccga gagacgcagg	4260
gagacgtaag agaggaaagt gtgagtggcc ggggggcctc cccccgtcag aagtcgcgca	4320
gtcgcgcccc taaaacgccc cctccgggcg gctagggcag gtgagcgct ccccgggcct	4380
ccccacgccc gcccctgcca caggccgtct aggtcgagca gatatttaca gaataaaaat	4440
gacaataact cgacgtcccc ggacggccac gcaatctgtt agtaatttag cgggatggga	4500
atttcctttc tagggcctgc cagtgaagcg cttttccaaa tttccacagc gggggaagcc	4560
tgcgatttta cataatgact tcagcatgcc gggctttctc gacacccctc cccggcccc	4620
ggccccgcc ccccgccct tttccagcag ggccgggctc cctccggaca cccgctgga	4680
ctcaggcgctc ccgtctggcc cgttcgcccc cgtttcccc gccagccca gcgccccct	4740
gccccggccc cggattcccc gttccgccc ctacgcccc atcccctccc cgtgcgcccc	4800
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ccgtgcgccc cccctcccc tgcgcgcccc gcctcttgcc cccctgcccc caggcgagcg	4980
gctgccgcgg cgcggggagg ggcgggcgct cggcgactcg tccccggggc cccgcgcggg	5040
cccgggcagc aggggcgtga tgtcacggca gggagggggc gcgggagccg ccgggccggc	5100
ggggaggcgg gggagggtgtt ttccagcttt aaaaaggcag gaggcagagc gcggccctgc	5160
gtcagagcga gactcagagg ctccgaactc gccggcggag tcgccgcgcc agatcccagc	5220
agcaggcgcc gggcacccgg gcgcgggcag ggctcggagc caccgcgag gtcctagggc	5280
cgcgccggg ccccgccacg gcgcacacg cccctcgatg	5320

<210> 816

<211> 361

<212> DNA

<213> Artificial Sequence

<220>

<223> NFATc3 promoter sequence

<400> 816

gcagccaggc aggggtgggcg gcgtagggg gcggggccgg gcgcgcggca gggcgcgaga	60
gcgcacccgc ggcggcggtg gcggcgactg tgggggggcg gcggggaaca ttggctaagc	120
cgacagtgga ggcttaggca ccgggtggcg gcggctgcgg ttcctggtgc tgctcggcgc	180
gcggccagct ttcggaacgg aacgctcggc gtcgcgggccc ccgcccggaa agtttgccgt	240
ggagtgcga cctcttggcc gcgcgggccc ggcatgaagc ggcgttgagg agctgctgcc	300
gccgcttgcc gctgccgccc ccgcccctg aggaggagct gcagcacccct gggccacgcc	360

<210> 817  
 <211> 430  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> NFATc2 promoter sequence 1

<400> 817  
 cagagagagg ctgcgttcag actggggcac tgccatcccc tccgcatcat ggggtctgtg 60  
 gaccaaggta actgactctc gatcccttcc agccttttcc gctcgtcctt cccggccctt 120  
 tcctgctgct cccgtcccgg gcagcacttt cagctcccgg cagaggtcgg tgcgggaggc 180  
 ctggggagccc cgctcgccct cggcgcacag gtagcggggc ccgcggaggg gcgcccgcgc 240  
 cccggccagg gaagggacac ttgggaaggc gactttggac aactttacgc gggggcaggg 300  
 aagtgtccca ggccgggatt ccctaggcca gtctgtcggg aggattttcc tctccacggg 360  
 acaccgggag ggattctcgc tactaaccgc tggctgttta accgtttcag cactcggctt 420  
 ttgacagcaa 430

<210> 818  
 <211> 64  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> NFATc2 promoter sequence 2

<400> 818  
 catcatgggg tctgtggacc aaggtaactg actctcgatc ctttcagcc ttttccgctc 60  
 gctc 64

<210> 819  
 <211> 12  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> NFATc1 response element consensus

<400> 819  
 cattttttcc at 12

<210> 820  
 <211> 8  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> NFATc1 response element consensus

<400> 820  
 tttttcca 8

<210> 821  
 <211> 119

<212> DNA  
<213> Artificial Sequence

<220>

<223> NFAT response elements contained within the Foxp3 enhancer region

<400> 821

acttgaaaat gagataaatg ttcacctatg ttggcttcta gtctctttta tggcttcatt 60

ttttccattt actatagagg ttaagagtgt gggctactgga gccagactgt ctgggacaa 119

<210> 822

<211> 65

<212> PRT

<213> Artificial Sequence

<220>

<223> N+20

<400> 822

Ser Asn Ile Lys Phe Arg Pro Gly Ser Val Val Val Gln Leu Thr Leu  
1 5 10 15

Ala Phe Arg Glu Gly Thr Ile Asn Val His Asp Val Glu Thr Gln Phe  
20 25 30

Asn Gln Tyr Lys Thr Glu Ala Ala Ser Arg Tyr Asn Leu Thr Ile Ser  
35 40 45

Asp Val Ser Val Ser Asp Val Pro Phe Pro Phe Ser Ala Gln Ser Gly  
50 55 60

Ala  
65

<210> 823

<211> 38

<212> PRT

<213> Artificial Sequence

<220>

<223> N+20/C27

<400> 823

Ser Asn Ile Lys Phe Arg Pro Gly Ser Val Val Val Gln Leu Thr Leu  
1 5 10 15

Ala Phe Arg Glu Gly Thr Ile Asn Val His Asp Val Glu Thr Gln Phe  
20 25 30

Asn Gln Tyr Lys Thr Glu  
35

<210> 824

<211> 45

<212> PRT

<213> Artificial Sequence



<220>  
<223> N+9/C9

<400> 824

Val Gln Leu Thr Leu Ala Phe Arg Glu Gly Thr Ile Asn Val His Asp  
1 5 10 15

Val Glu Thr Gln Phe Asn Gln Tyr Lys Thr Glu Ala Ala Ser Arg Tyr  
20 25 30

Asn Leu Thr Ile Ser Asp Val Ser Val Ser Asp Val Pro  
35 40 45

<210> 825  
<211> 35  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> C10

<400> 825

Gly Thr Ile Asn Val His Asp Val Glu Thr Gln Phe Asn Gln Tyr Lys  
1 5 10 15

Thr Glu Ala Ala Ser Arg Tyr Asn Leu Thr Ile Ser Asp Val Ser Val  
20 25 30

Ser Asp Val  
35

<210> 826  
<211> 885  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> NME7AB

<400> 826

atggaaaaaa cgctggccct gattaaaccg gatgcaatct ccaaagctgg cgaaattatc	60
gaaattatca acaaagcggg ttccaccatc acgaaactga aaatgatgat gctgagccgt	120
aaagaagccc tggattttca tgtcgaccac cagtctcgcc cgtttttcaa tgaactgatt	180
caattcatca ccacgggtcc gattatcgca atggaaattc tgcgtgatga cgctatctgc	240
gaatggaaac gcctgctggg cccggcaaac tcaggtgttg cgcgtaccga tgccagtga	300
tccattcgcg ctctgtttgg caccgatggg atccgtaatg cagcacatgg tccggactca	360
ttcgcacgag cagctcgtga aatggaactg tttttcccga gctctggcgg ttgcgggtccg	420
gcaaacaccg ccaaatttac caattgtacg tgctgtattg tcaaaccgca cgcagtgtca	480
gaaggcctgc tgggtaaaat tctgatggca atccgtgatg ctggctttga aatctcggcc	540
atgcagatgt tcaacatgga ccgcgttaac gtcgaagaat tctacgaagt ttacaaaggc	600
gtggttaccg aatatcacga tatggttacg gaaatgtact ccggtccgtg cgtcgcgatg	660

gaaattcagc aaaacaatgc caccaaaacg tttcgtgaat tctgtggtcc ggcagatccg	720
gaaatcgcac gtcattctgcg tccgggtacc ctgcgcgcaa tttttggtaa aacgaaaatc	780
cagaacgctg tgcactgtac cgatctgccg gaagacggtc tgctggaagt tcaatacttt	840
ttcaaaattc tggataatct cgagcaccac caccaccacc actga	885

<210> 827  
 <211> 294  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> NME7AB

<400> 827

Met	Glu	Lys	Thr	Leu	Ala	Leu	Ile	Lys	Pro	Asp	Ala	Ile	Ser	Lys	Ala
1				5				10						15	

Gly	Glu	Ile	Ile	Glu	Ile	Ile	Asn	Lys	Ala	Gly	Phe	Thr	Ile	Thr	Lys
		20					25						30		

Leu	Lys	Met	Met	Met	Leu	Ser	Arg	Lys	Glu	Ala	Leu	Asp	Phe	His	Val
		35					40					45			

Asp	His	Gln	Ser	Arg	Pro	Phe	Phe	Asn	Glu	Leu	Ile	Gln	Phe	Ile	Thr
	50					55					60				

Thr	Gly	Pro	Ile	Ile	Ala	Met	Glu	Ile	Leu	Arg	Asp	Asp	Ala	Ile	Cys
65					70					75					80

Glu	Trp	Lys	Arg	Leu	Leu	Gly	Pro	Ala	Asn	Ser	Gly	Val	Ala	Arg	Thr
			85						90					95	

Asp	Ala	Ser	Glu	Ser	Ile	Arg	Ala	Leu	Phe	Gly	Thr	Asp	Gly	Ile	Arg
		100						105					110		

Asn	Ala	Ala	His	Gly	Pro	Asp	Ser	Phe	Ala	Ser	Ala	Ala	Arg	Glu	Met
		115					120						125		

Glu	Leu	Phe	Phe	Pro	Ser	Ser	Gly	Gly	Cys	Gly	Pro	Ala	Asn	Thr	Ala
	130					135					140				

Lys	Phe	Thr	Asn	Cys	Thr	Cys	Cys	Ile	Val	Lys	Pro	His	Ala	Val	Ser
145					150					155					160

Glu	Gly	Leu	Leu	Gly	Lys	Ile	Leu	Met	Ala	Ile	Arg	Asp	Ala	Gly	Phe
				165					170					175	

Glu	Ile	Ser	Ala	Met	Gln	Met	Phe	Asn	Met	Asp	Arg	Val	Asn	Val	Glu
			180					185					190		

Glu	Phe	Tyr	Glu	Val	Tyr	Lys	Gly	Val	Val	Thr	Glu	Tyr	His	Asp	Met
-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----

195

200

205

Val Thr Glu Met Tyr Ser Gly Pro Cys Val Ala Met Glu Ile Gln Gln  
 210 215 220

Asn Asn Ala Thr Lys Thr Phe Arg Glu Phe Cys Gly Pro Ala Asp Pro  
 225 230 235 240

Glu Ile Ala Arg His Leu Arg Pro Gly Thr Leu Arg Ala Ile Phe Gly  
 245 250 255

Lys Thr Lys Ile Gln Asn Ala Val His Cys Thr Asp Leu Pro Glu Asp  
 260 265 270

Gly Leu Leu Glu Val Gln Tyr Phe Phe Lys Ile Leu Asp Asn Leu Glu  
 275 280 285

His His His His His His  
 290

<210> 828  
 <211> 783  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Human NME7 x1

<400> 828  
 atgatgatgc tttcaaggaa agaagcattg gattttcatg tagatcacca gtcaagaccc 60  
 tttttcaatg agctgatcca gtttattaca actggtccta ttattgccat ggagatttta 120  
 agagatgatg ctatatgtga atggaaaaga ctgctgggac ctgcaaactc tggagtggca 180  
 cgcacagatg cttctgaaag cattagagcc ctctttggaa cagatggcat aagaaatgca 240  
 gcgcattggcc ctgattcttt tgcttctgcg gccagagaaa tggagtgtgt ttttccttca 300  
 agtggagggtt gtgggccggc aaacactgct aaatttacta attgtacctg ttgcattgtt 360  
 aaaccccatg ctgtcagtga aggactgttg ggaaagatcc tgatggctat ccgagatgca 420  
 ggttttgaaa tctcagctat gcagatgttc aatatggatc gggttaatgt tgaggaattc 480  
 tatgaagttt ataaaggagt agtgaccgaa tatcatgaca tggtgacaga aatgtattct 540  
 ggcccttgtg tagcaatgga gattcaacag aataatgcta caaagacatt tcgagaattt 600  
 tgtggacctg ctgatcctga aattgcccgg catttacgcc ctggaactct cagagcaatc 660  
 tttggtaaaa ctaagatcca gaatgctgtt cactgtactg atctgccaga ggatggccta 720  
 ttagaggttc aatacttctt caagatcttg gataatctcg agcaccacca ccaccaccac 780  
 tga 783

<210> 829  
 <211> 260  
 <212> PRT  
 <213> Artificial Sequence

<220>

<223> Human NME7 x1

<400> 829

Met Met Met Leu Ser Arg Lys Glu Ala Leu Asp Phe His Val Asp His  
1 5 10 15

Gln Ser Arg Pro Phe Phe Asn Glu Leu Ile Gln Phe Ile Thr Thr Gly  
20 25 30

Pro Ile Ile Ala Met Glu Ile Leu Arg Asp Asp Ala Ile Cys Glu Trp  
35 40 45

Lys Arg Leu Leu Gly Pro Ala Asn Ser Gly Val Ala Arg Thr Asp Ala  
50 55 60

Ser Glu Ser Ile Arg Ala Leu Phe Gly Thr Asp Gly Ile Arg Asn Ala  
65 70 75 80

Ala His Gly Pro Asp Ser Phe Ala Ser Ala Ala Arg Glu Met Glu Leu  
85 90 95

Phe Phe Pro Ser Ser Gly Gly Cys Gly Pro Ala Asn Thr Ala Lys Phe  
100 105 110

Thr Asn Cys Thr Cys Cys Ile Val Lys Pro His Ala Val Ser Glu Gly  
115 120 125

Leu Leu Gly Lys Ile Leu Met Ala Ile Arg Asp Ala Gly Phe Glu Ile  
130 135 140

Ser Ala Met Gln Met Phe Asn Met Asp Arg Val Asn Val Glu Glu Phe  
145 150 155 160

Tyr Glu Val Tyr Lys Gly Val Val Thr Glu Tyr His Asp Met Val Thr  
165 170 175

Glu Met Tyr Ser Gly Pro Cys Val Ala Met Glu Ile Gln Gln Asn Asn  
180 185 190

Ala Thr Lys Thr Phe Arg Glu Phe Cys Gly Pro Ala Asp Pro Glu Ile  
195 200 205

Ala Arg His Leu Arg Pro Gly Thr Leu Arg Ala Ile Phe Gly Lys Thr  
210 215 220

Lys Ile Gln Asn Ala Val His Cys Thr Asp Leu Pro Glu Asp Gly Leu  
225 230 235 240

Leu Glu Val Gln Tyr Phe Phe Lys Ile Leu Asp Asn Leu Glu His His  
245 250 255

His His His His  
260

<210> 830  
<211> 426  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse Antibody 17H6 Heavy chain: DNA sequence - Signal sequence  
FR1CDR1FR2CDR2FR3CDR3FR4

<400> 830  
atgaagttgt ggctgaactg gattttcctt gtaacacttt taaatggtat ccagtgtgag 60  
gtgaagctgg tggagtctgg aggaggcttg gtacagcctg ggggttctct gagactctcc 120  
tgtgcaactt ctgggttcac cttcactgat tactacatga gctgggtccg ccagcctcca 180  
agaaaggcac ttgagtgggtt ggggttttatt agaaacaaag ctaatggta cacagcagag 240  
tacagtgcgt ctgtgaaggg tcggttcacc atctccagag atgtttcca aaacctctc 300  
tatcttcaaa tgaacatcct gagagctgag gacagtgcc cttattactg tgcaaaagat 360  
tactacggtg gtaaccctgc ctggtttgct tactggggcc aagggactct ggtcactgtc 420  
tctgca 426

<210> 831  
<211> 142  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse Antibody 17H6 Heavy chain: Amino acid sequence - Signal  
peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 831

Met Lys Leu Trp Leu Asn Trp Ile Phe Leu Val Thr Leu Leu Asn Gly  
1 5 10 15

Ile Gln Cys Glu Val Lys Leu Val Glu Ser Gly Gly Gly Leu Val Gln  
20 25 30

Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Thr Ser Gly Phe Thr Phe  
35 40 45

Thr Asp Tyr Tyr Met Ser Trp Val Arg Gln Pro Pro Arg Lys Ala Leu  
50 55 60

Glu Trp Leu Gly Phe Ile Arg Asn Lys Ala Asn Gly Tyr Thr Ala Glu  
65 70 75 80

Tyr Ser Ala Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Val Ser  
85 90 95

Gln Asn Leu Leu Tyr Leu Gln Met Asn Ile Leu Arg Ala Glu Asp Ser  
100 105 110

Ala Thr Tyr Tyr Cys Ala Lys Asp Tyr Tyr Gly Ser Asn Pro Ala Trp  
115 120 125

Phe Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
130 135 140

<210> 832  
<211> 90  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse 17H6 heavy chain variable framework 1 (FW1) sequence

<400> 832  
gaggatgaagc tgggtggagtc tggaggaggc ttggtacagc ctgggggttc tctgagactc 60  
tcctgtgcaa cttctgggtt caccttcact 90

<210> 833  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 17H6 heavy chain variable framework 1 (FW1) sequence

<400> 833

Glu Val Lys Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Thr Ser Gly Phe Thr Phe Thr  
20 25 30

<210> 834  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse 17H6 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 834  
gattactaca tgagc 15

<210> 835  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 17H6 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 835

Asp Tyr Tyr Met Ser  
1 5

<210> 836  
 <211> 90  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 heavy chain variable framework 2 (FW2) sequence  
  
 <400> 836  
 gaggtgaagc tgggtggagtc tggaggaggc ttggtacagc ctggggggtc tctgagactc 60  
 tcctgtgcaa cttctggggtt caccttcact 90

<210> 837  
 <211> 14  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 heavy chain variable framework 2 (FW2) sequence  
  
 <400> 837  
  
 Trp Val Arg Gln Pro Pro Arg Lys Ala Leu Glu Trp Leu Gly  
 1 5 10

<210> 838  
 <211> 57  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 838  
 tttattagaa acaaagctaa tggttacaca gcagagtaca gtgcgtctgt gaagggt 57

<210> 839  
 <211> 19  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 839  
  
 Phe Ile Arg Asn Lys Ala Asn Gly Tyr Thr Ala Glu Tyr Ser Ala Ser  
 1 5 10 15

Val Lys Gly

<210> 840  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 heavy chain variable framework 3 (FW3) sequence

<400> 840  
 cgggtcacca tctccagaga tgtttcccaa aacctctct atcttcaa gaacatcctg 60  
 agagctgagg acagtgccac ttattactgt gcaaaa 96

<210> 841  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse 17H6 heavy chain variable framework 3 (FW3) sequence

<400> 841  
 Arg Phe Thr Ile Ser Arg Asp Val Ser Gln Asn Leu Leu Tyr Leu Gln  
 1 5 10 15  
 Met Asn Ile Leu Arg Ala Glu Asp Ser Ala Thr Tyr Tyr Cys Ala Lys  
 20 25 30

<210> 842  
 <211> 36  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse 17H6 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 842  
 gattactacg gtagtaaccc tgcctggttt gcttac 36

<210> 843  
 <211> 33  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse 17H6 heavy chain variable framework 4 (FW4) sequence

<400> 843  
 tggggccaag ggactctggt cactgtctct gca 33

<210> 844  
 <211> 393  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse Antibody 17H6 Light chain: DNA sequence - Signal sequence  
 FR1CDR1FR2CDR2FR3CDR3FR4

<400> 844  
 atgaagttgc ctgtgaggct gttggtgctg atgttctgga ttcttgcttc caacagtgat 60  
 attttgatga cccagactcc actctccctg cctgtcagtc ttggagatca agcctccatc 120  
 tcttgcatga ctagtcagag cattgtacat agtagtgga acacctttt agaatggtac 180  
 ctgcagaaac ctggccagtc tccaaagctc ctgatctaca aagtttccaa ccgattttct 240  
 ggggtcccag acaggttcag tggcagtgga tcagggatag atttcacact caagatcagc 300



agagtggagg ctgaggatct gggagtttat tactgctttc aaggttcaca tgttcctttc 360

acgttcggct cggggacaaa gttggaaata aaa 393

<210> 845

<211> 131

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse Antibody 17H6 Light chain: Amino acid sequence - Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 845

Met Lys Leu Pro Val Arg Leu Leu Val Leu Met Phe Trp Ile Pro Ala  
1 5 10 15

Ser Asn Ser Asp Ile Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val  
20 25 30

Ser Leu Gly Asp Gln Ala Ser Ile Ser Cys Arg Ser Ser Gln Ser Ile  
35 40 45

Val His Ser Ser Gly Asn Thr Phe Leu Glu Trp Tyr Leu Gln Lys Pro  
50 55 60

Gly Gln Ser Pro Lys Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser  
65 70 75 80

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Ile Asp Phe Thr  
85 90 95

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
100 105 110

Phe Gln Gly Ser His Val Pro Phe Thr Phe Gly Ser Gly Thr Lys Leu  
115 120 125

Glu Ile Lys  
130

<210> 846

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 17H6 light chain variable framework 1 (FW1) sequence

<400> 846

gatattttga tgaccagac tccactctcc ctgcctgtca gtcttgaga tcaagcctcc 60

atctcttgcc 69

<210> 847

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 17H6 light chain variable framework 1 (FW1) sequence

<400> 847

Asp Ile Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
1 5 10 15

Asp Gln Ala Ser Ile Ser Cys  
20

<210> 848

<211> 48

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 17H6 light chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 848

agatctagtc agagcattgt acatagtagt ggaaacacct ttttagaa 48

<210> 849

<211> 16

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 17H6 light chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 849

Arg Ser Ser Gln Ser Ile Val His Ser Ser Gly Asn Thr Phe Leu Glu  
1 5 10 15

<210> 850

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 17H6 light chain variable framework 2 (FW2) sequence

<400> 850

tggtacctgc agaaacctgg ccagtctcca aagctcctga tctac 45

<210> 851

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 17H6 light chain variable framework 2 (FW2) sequence

<400> 851

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 852  
 <211> 21  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 light chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 852  
 aaagtttcca accgattttc t 21

<210> 853  
 <211> 7  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 light chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 853

Lys Val Ser Asn Arg Phe Ser  
 1 5

<210> 854  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 light chain variable framework 3 (FW3) sequence  
  
 <400> 854  
 ggggtcccag acaggttcag tggcagtgga tcagggatag atttcacact caagatcagc 60  
 agagtggagg ctgaggatct gggagtttat tactgc 96

<210> 855  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 light chain variable framework 3 (FW3) sequence  
  
 <400> 855

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Ile Asp Phe Thr  
 1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
 20 25 30

<210> 856  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 17H6 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 856  
tttcaagggtt cacatgttcc tttcacg 27

<210> 857  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 17H6 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 857  
Phe Gln Gly Ser His Val Pro Phe Thr  
1 5

<210> 858  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse 17H6 light chain variable framework 4 (FW4) sequence

<400> 858  
ttcggctcgg ggacaaagtt ggaaataaaa 30

<210> 859  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 17H6 light chain variable framework 4 (FW4) sequence

<400> 859  
Phe Gly Ser Gly Thr Lys Leu Glu Ile Lys  
1 5 10

<210> 860  
<211> 408  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 39H5 Heavy chain: DNA sequence - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 860  
atggccttggg tgtggacctt gctattcctg atggcagctg cccaaagtgc ccaagcacag 60  
atccagttgg tgcagtctgg acctgagctg aagaagcctg gagagacagt caagatctcc 120  
tgcaaggctt ctgggtatac cttcaciaaac tatggaatga actgggtgaa gcaggctcca 180  
ggaaagggtt taaagtggat gggctggata aacacctaca ctggagagcc aacatatgtt 240  
ggtgacttca agggacgggt tgccttctct ttggagacct ctgccagcac tgcctatttg 300  
cagatcaaca acctcaaaaa tgaggacacg gctacatatt tttgtgttag aggtatccac 360  
ggctacgtgg actactgggg ccaaggcacc actctcacag tctcctca 408

<210> 861  
 <211> 136  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 Heavy chain: Amino acid sequence - Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 861

Met Ala Trp Val Trp Thr Leu Leu Phe Leu Met Ala Ala Ala Gln Ser  
 1 5 10 15

Ala Gln Ala Gln Ile Gln Leu Val Gln Ser Gly Pro Glu Leu Lys Lys  
 20 25 30

Pro Gly Glu Thr Val Lys Ile Ser Cys Lys Ala Ser Gly Tyr Thr Phe  
 35 40 45

Thr Asn Tyr Gly Met Asn Trp Val Lys Gln Ala Pro Gly Lys Gly Leu  
 50 55 60

Lys Trp Met Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Val  
 65 70 75 80

Gly Asp Phe Lys Gly Arg Phe Ala Phe Ser Leu Glu Thr Ser Ala Ser  
 85 90 95

Thr Ala Tyr Leu Gln Ile Asn Asn Leu Lys Asn Glu Asp Thr Ala Thr  
 100 105 110

Tyr Phe Cys Val Arg Gly Ile His Gly Tyr Val Asp Tyr Trp Gly Gln  
 115 120 125

Gly Thr Thr Leu Thr Val Ser Ser  
 130 135

<210> 862  
 <211> 90  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 39H5 heavy chain variable framework 1 (FW1) sequence

<400> 862

cagatccagt tgggtgcagtc tggacctgag ctgaagaagc ctggagagac agtcaagatc 60

tcctgcaagg cttctgggta taccttcaca 90

<210> 863  
 <211> 30  
 <212> PRT  
 <213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable framework 1 (FW1) sequence

<400> 863

Gln Ile Gln Leu Val Gln Ser Gly Pro Glu Leu Lys Lys Pro Gly Glu  
1 5 10 15

Thr Val Lys Ile Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr  
20 25 30

<210> 864

<211> 15

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 864

aactatggaa tgaac

15

<210> 865

<211> 5

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 865

Asn Tyr Gly Met Asn  
1 5

<210> 866

<211> 42

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable framework 2 (FW2) sequence

<400> 866

tgggtgaagc aggctccagg aaagggttta aagtggatgg gc

42

<210> 867

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable framework 2 (FW2) sequence

<400> 867

Trp Val Lys Gln Ala Pro Gly Lys Gly Leu Lys Trp Met Gly  
1 5 10

<210> 868  
 <211> 51  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 heavy chain variable complementarity  
 determining regions 2 (CDR2) sequence  
  
 <400> 868  
 tggataaaca cctacactgg agagccaaca tatgttggtg acttcaaggg a 51

<210> 869  
 <211> 17  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 heavy chain variable complementarity  
 determining regions 2 (CDR2) sequence

<400> 869  
  
 Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Val Gly Asp Phe Lys  
 1 5 10 15

Gly

<210> 870  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 heavy chain variable framework 3 (FW3)  
 sequence  
  
 <400> 870  
 cgggtttgcct tctcttttga gacctctgcc agcactgcct atttgcagat caacaacctc 60  
  
 aaaaatgagg acacggctac atatTTTTgt gttaga 96

<210> 871  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 heavy chain variable framework 3 (FW3)  
 sequence

<400> 871  
  
 Arg Phe Ala Phe Ser Leu Glu Thr Ser Ala Ser Thr Ala Tyr Leu Gln  
 1 5 10 15

Ile Asn Asn Leu Lys Asn Glu Asp Thr Ala Thr Tyr Phe Cys Val Arg  
 20 25 30

<210> 872  
 <211> 24  
 <212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 872

ggtatccacg gctacgtgga ctac

24

<210> 873

<211> 8

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 873

Gly Ile His Gly Tyr Val Asp Tyr

1 5

<210> 874

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable framework 4 (FW4) sequence

<400> 874

tggggccaag gcaccactct cacagtctcc tca

33

<210> 875

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable framework 4 (FW4) sequence

<400> 875

Trp Gly Gln Gly Thr Thr Leu Thr Val Ser Ser

1 5 10

<210> 876

<211> 393

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 Light chain: DNA sequence - Signal sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 876

atgaagttgc ctgttaggct gttggtgctg atgttctgga ttcctgcttc cagcagtgat

60

gttttgatga cccaaactcc actctccctg cctgtcagtc ttggagatca agcctccatc

120

tcttcagat ctagtcagag cattgtacat agaaatggaa acacctatctt agaatggtag

180



ctgcagaaac caggccagtc tccaaagctc ctgatctaca aagtttccaa ccgattttct	240
ggggtcccag acaggttcag tggcagtgga tcagggacag atttcacact caagatcagc	300
agagtggagg ctgaggatct gggagtttat tactgctttc aaggttcaca tcttcgtgg	360
acgttcggtg gaggcaccaa gctggaaatc aaa	393

<210> 877  
 <211> 131  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 39H5 Light chain: Amino acid sequence - Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 877

Met	Lys	Leu	Pro	Val	Arg	Leu	Leu	Val	Leu	Met	Phe	Trp	Ile	Pro	Ala
1				5					10					15	

Ser	Ser	Ser	Asp	Val	Leu	Met	Thr	Gln	Thr	Pro	Leu	Ser	Leu	Pro	Val
			20					25					30		

Ser	Leu	Gly	Asp	Gln	Ala	Ser	Ile	Ser	Cys	Arg	Ser	Ser	Gln	Ser	Ile
		35					40					45			

Val	His	Arg	Asn	Gly	Asn	Thr	Tyr	Leu	Glu	Trp	Tyr	Leu	Gln	Lys	Pro
	50					55					60				

Gly	Gln	Ser	Pro	Lys	Leu	Leu	Ile	Tyr	Lys	Val	Ser	Asn	Arg	Phe	Ser
65					70					75					80

Gly	Val	Pro	Asp	Arg	Phe	Ser	Gly	Ser	Gly	Ser	Gly	Thr	Asp	Phe	Thr
				85					90					95	

Leu	Lys	Ile	Ser	Arg	Val	Glu	Ala	Glu	Asp	Leu	Gly	Val	Tyr	Tyr	Cys
			100					105					110		

Phe	Gln	Gly	Ser	His	Leu	Pro	Trp	Thr	Phe	Gly	Gly	Gly	Thr	Lys	Leu
		115					120					125			

Glu	Ile	Lys
		130

<210> 878  
 <211> 69  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 39H5 light chain variable framework 1 (FW1) sequence

<400> 878

gatgttttga tgacccaaac tccactctcc ctgcctgtca gtcttgaga tcaagcctcc	60
atctcttgc	69

<210> 879  
 <211> 23  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 light chain variable framework 1 (FW1) sequence

<400> 879  
  
 Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
 1 5 10 15

Asp Gln Ala Ser Ile Ser Cys  
 20

<210> 880  
 <211> 48  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 880  
 agatctagtc agagcattgt acatagaaat ggaaacacct atttagaa 48

<210> 881  
 <211> 16  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 881  
  
 Arg Ser Ser Gln Ser Ile Val His Arg Asn Gly Asn Thr Tyr Leu Glu  
 1 5 10 15

<210> 882  
 <211> 45  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 light chain variable framework 2 (FW2) sequence

<400> 882  
 tggtagctgc agaaaccagg ccagttctcca aagctcctga tctac 45

<210> 883  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 light chain variable framework 2 (FW2)

sequence

<400> 883

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 884

<211> 21

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 884

aaagtttcca accgattttc t

21

<210> 885

<211> 7

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 885

Lys Val Ser Asn Arg Phe Ser  
1 5

<210> 886

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 light chain variable framework 3 (FW3)  
sequence

<400> 886

gggggtcccg acaggttcag tggcagtgga tcagggacag atttcacact caagatcagc

60

agagtggagg ctgaggatct gggagtttat tactgc

96

<210> 887

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 light chain variable framework 3 (FW3)  
sequence

<400> 887

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
20 25 30

<210> 888  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 light chain variable complementarity  
 determining regions 3 (CDR3) sequence  
  
 <400> 888  
 tttcaaggtt cacatcttcc gtggacg 27

<210> 889  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 light chain variable complementarity  
 determining regions 3 (CDR3) sequence  
  
 <400> 889  
  
 Phe Gln Gly Ser His Leu Pro Trp Thr  
 1 5

<210> 890  
 <211> 30  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 light chain variable framework 4 (FW4)  
 sequence  
  
 <400> 890  
 ttcggtggag gcaccaagct ggaaatcaaa 30

<210> 891  
 <211> 10  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 39H5 light chain variable framework 4 (FW4)  
 sequence  
  
 <400> 891  
  
 Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
 1 5 10

<210> 892  
 <211> 411  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Antibody 3C5 Heavy chain: DNA sequence - Signal  
 sequenceFR1CDR1FR2CDR2FR3CDR3FR4  
  
 <400> 892  
 atggcttggg tgtggacctt gctgttcctg atggcagctg cccaaagtgc ccaagcacag 60

atccagttgg tgcagtctgg acctgagctg aagaagcctg gagagacagt caagatctcc 120  
 tgcaaggctt ctgggtatac cttcacaac tatggaatga actgggtgaa gcaggctcca 180  
 ggaaagggtt taaagtggat gggctggata aacacctaca ctggaaagcc aacatatgct 240  
 gatgacttca agggacggtt tgccttctct ttggagacct ctgccagcac tgcctatttg 300  
 cagatcaaca acctcaaaaa tgaggacacg gctacatatt tctgtgcaag aggggggacta 360  
 gatggttact acggctactg gggccaaggc accactctca cagtctcctc a 411

<210> 893  
 <211> 137  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Antibody 3C5 Heavy chain: Amino acid sequence - Signal  
 peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 893

Met Ala Trp Val Trp Thr Leu Leu Phe Leu Met Ala Ala Ala Gln Ser  
 1 5 10 15

Ala Gln Ala Gln Ile Gln Leu Val Gln Ser Gly Pro Glu Leu Lys Lys  
 20 25 30

Pro Gly Glu Thr Val Lys Ile Ser Cys Lys Ala Ser Gly Tyr Thr Phe  
 35 40 45

Thr Asn Tyr Gly Met Asn Trp Val Lys Gln Ala Pro Gly Lys Gly Leu  
 50 55 60

Lys Trp Met Gly Trp Ile Asn Thr Tyr Thr Gly Lys Pro Thr Tyr Ala  
 65 70 75 80

Asp Asp Phe Lys Gly Arg Phe Ala Phe Ser Leu Glu Thr Ser Ala Ser  
 85 90 95

Thr Ala Tyr Leu Gln Ile Asn Asn Leu Lys Asn Glu Asp Thr Ala Thr  
 100 105 110

Tyr Phe Cys Ala Arg Gly Gly Leu Asp Gly Tyr Tyr Gly Tyr Trp Gly  
 115 120 125

Gln Gly Thr Thr Leu Thr Val Ser Ser  
 130 135

<210> 894  
 <211> 90  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 3C5 heavy chain variable framework 1 (FW1)  
 sequence

<400> 894  
cagatccagt tgggtgcagtc tggacctgag ctgaagaagc ctggagagac agtcaagatc 60  
tcctgcaagg cttctgggta taccttcaca 90

<210> 895  
<211> 30  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 3C5 heavy chain variable framework 1 (FW1)  
sequence

<400> 895  
Gln Ile Gln Leu Val Gln Ser Gly Pro Glu Leu Lys Lys Pro Gly Glu  
1 5 10 15

Thr Val Lys Ile Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr  
20 25 30

<210> 896  
<211> 15  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 3C5 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 896  
aactatggaa tgaac 15

<210> 897  
<211> 5  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 3C5 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 897  
Asn Tyr Gly Met Asn  
1 5

<210> 898  
<211> 42  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 3C5 heavy chain variable framework 2 (FW2)  
sequence

<400> 898  
tgggtgaagc aggctccagg aaagggttta aagtggatgg gc 42

<210> 899  
<211> 14  
<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 heavy chain variable framework 2 (FW2) sequence

<400> 899

Trp Val Lys Gln Ala Pro Gly Lys Gly Leu Lys Trp Met Gly  
1 5 10

<210> 900

<211> 51

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 900

tggataaaca cctacactgg aaagccaaca tatgctgatg acttcaaggg a 51

<210> 901

<211> 17

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 901

Trp Ile Asn Thr Tyr Thr Gly Lys Pro Thr Tyr Ala Asp Asp Phe Lys  
1 5 10 15

Gly

<210> 902

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 heavy chain variable framework 3 (FW3) sequence

<400> 902

cggtttgctt tctcttttga gacctctgcc agcactgcct atttcagat caacaacctc 60

aaaaatgagg acacggctac atatttctgt gcaaga 96

<210> 903

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 heavy chain variable framework 3 (FW3) sequence

<400> 903

Arg Phe Ala Phe Ser Leu Glu Thr Ser Ala Ser Thr Ala Tyr Leu Gln  
1 5 10 15

Ile Asn Asn Leu Lys Asn Glu Asp Thr Ala Thr Tyr Phe Cys Ala Arg  
20 25 30

<210> 904  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 3C5 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 904  
gggggactag atggttacta cggctac

27

<210> 905  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 3C5 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 905

Gly Gly Leu Asp Gly Tyr Tyr Gly Tyr  
1 5

<210> 906  
<211> 33  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 3C5 heavy chain variable framework 4 (FW4)  
sequence

<400> 906  
tggggccaag gcaccactct cacagtctcc tca

33

<210> 907  
<211> 11  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 3C5 heavy chain variable framework 4 (FW4)  
sequence

<400> 907

Trp Gly Gln Gly Thr Thr Leu Thr Val Ser Ser  
1 5 10

<210> 908  
<211> 393  
<212> DNA  
<213> Artificial Sequence



<220>

<223> Antibody 3C5 Light chain: DNA sequence (393 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 908

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atgagtcctg cccagttcct gtttctgcta gtgctctcga ttcaggaaac caacggtgat      60
gttgatgatgg ctcagacccc actcactttg tcggttacca ttggacaacc agcctccatc      120
tcttgcaaat caagtcagag cctcttacat agtaaaggaa agacatattt gaattgggta      180
ttacagaggc caggccagtc tccaaagctc ctaatctatc tgggtgtctaa actggaatct      240
ggagtcacctg acaggttcag tggcagtgga tcagggacag atttcacact gaaaatcagc      300
agagtggagg ctgaagattt gggagtttat tactgcttgc aaactacaca ttttccgtgg      360
acgttcggtg gaggcaccaa gctggaaatc aaa                                  393
```

<210> 909

<211> 131

<212> PRT

<213> Artificial Sequence

<220>

<223> Antibody 3C5 Light chain: Amino acid sequence (131 aa) - Signal  
peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 909

```
Met Ser Pro Ala Gln Phe Leu Phe Leu Leu Val Leu Ser Ile Gln Glu
1          5          10          15
```

```
Thr Asn Gly Asp Val Val Met Ala Gln Thr Pro Leu Thr Leu Ser Val
          20          25          30
```

```
Thr Ile Gly Gln Pro Ala Ser Ile Ser Cys Lys Ser Ser Gln Ser Leu
          35          40          45
```

```
Leu His Ser Lys Gly Lys Thr Tyr Leu Asn Trp Leu Leu Gln Arg Pro
          50          55          60
```

```
Gly Gln Ser Pro Lys Leu Leu Ile Tyr Leu Val Ser Lys Leu Glu Ser
          65          70          75          80
```

```
Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr
          85          90          95
```

```
Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys
          100         105         110
```

```
Leu Gln Thr Thr His Phe Pro Trp Thr Phe Gly Gly Gly Thr Lys Leu
          115         120         125
```

```
Glu Ile Lys
          130
```

<210> 910

<211> 69  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 3C5 light chain variable framework 1 (FW1) sequence

<400> 910  
 gatgttgatga tggctcagac cccactcact ttgtcgggta ccattggaca accagcctcc 60  
 atctcttgc 69

<210> 911  
 <211> 23  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 3C5 light chain variable framework 1 (FW1) sequence

<400> 911  
 Asp Val Val Met Ala Gln Thr Pro Leu Thr Leu Ser Val Thr Ile Gly  
 1 5 10 15  
 Gln Pro Ala Ser Ile Ser Cys  
 20

<210> 912  
 <211> 48  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 39H5 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 912  
 aaatcaagtc agagcctctt acatagtaaa ggaaagacat atttgaat 48

<210> 913  
 <211> 16  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 39H5 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 913  
 Lys Ser Ser Gln Ser Leu Leu His Ser Lys Gly Lys Thr Tyr Leu Asn  
 1 5 10 15

<210> 914  
 <211> 45  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 3C5 light chain variable framework 2 (FW2) sequence

<400> 914  
tggttattac agaggccagg ccagtctcca aagctcctaa tctat 45

<210> 915  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 3C5 light chain variable framework 2 (FW2)  
sequence

<400> 915  
Trp Leu Leu Gln Arg Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 916  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 3C5 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 916  
ctggtgtcta aactggaatc t 21

<210> 917  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 3C5 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 917  
Leu Val Ser Lys Leu Glu Ser  
1 5

<210> 918  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 3C5 light chain variable framework 3 (FW3)  
sequence

<400> 918  
ggagtccttg acaggttcag tggcagtgga tcagggacag atttcacact gaaaatcagc 60  
agagtggagg ctgaagattt gggagtttat tactgc 96

<210> 919  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 light chain variable framework 3 (FW3) sequence

<400> 919

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
20 25 30

<210> 920

<211> 27

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 920

ttgcaaaacta cacattttcc gtggacg

27

<210> 921

<211> 9

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 921

Leu Gln Thr Thr His Phe Pro Trp Thr  
1 5

<210> 922

<211> 30

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 light chain variable framework 4 (FW4) sequence

<400> 922

ttcgggtggag gcaccaagct ggaaatcaaa

30

<210> 923

<211> 10

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 3C5 light chain variable framework 4 (FW4) sequence

<400> 923

Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
1 5 10

<210> 924  
<211> 420  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 Heavy chain: DNA sequence (420 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 924  
atgaagtgtg ggctgaactg gattttcctt gtaacacttt taaatggat ccagtgtgag 60  
gtggagctgg tggagtctgg aggaggcttg gtacagcctg ggggttctct gagactctcc 120  
tgtgcaactt ctgggttcac cttcactgat cactacatga gctgggtccg ccagcctcca 180  
ggaaaggcac ttgagtgggtt gggatttatt agaaacaaag ctaatggta cacaacagag 240  
tacagtgcac ctgtgaaggg tcggttcacc atctccagag ataattccca aagcatcctc 300  
tatcttcaaa tgaaaacctt gagaactgag gacagtgcc cttattactg tgcaagacct 360  
tctgactggg actcctgggtt tgcttactgg ggccaaggga ctctggtcac tgtctctgca 420

<210> 925  
<211> 140  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 Heavy chain: Amino acid sequence (140 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 925  
Met Lys Leu Trp Leu Asn Trp Ile Phe Leu Val Thr Leu Leu Asn Gly  
1 5 10 15  
Ile Gln Cys Glu Val Glu Leu Val Glu Ser Gly Gly Gly Leu Val Gln  
20 25 30  
Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Thr Ser Gly Phe Thr Phe  
35 40 45  
Thr Asp His Tyr Met Ser Trp Val Arg Gln Pro Pro Gly Lys Ala Leu  
50 55 60  
Glu Trp Leu Gly Phe Ile Arg Asn Lys Ala Asn Gly Tyr Thr Thr Glu  
65 70 75 80  
Tyr Ser Ala Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ser  
85 90 95  
Gln Ser Ile Leu Tyr Leu Gln Met Lys Thr Leu Arg Thr Glu Asp Ser  
100 105 110  
Ala Thr Tyr Tyr Cys Ala Arg Pro Ser Asp Trp Asp Ser Trp Phe Ala  
115 120 125  
Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala

<210> 926  
 <211> 90  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 heavy chain variable framework 1 (FW1) sequence

<400> 926  
 gaggtggagc tgggtggagtc tggaggaggc ttggtacagc ctgggggttc tctgagactc 60  
 tcctgtgcaa cttctgggtt caccttcact 90

<210> 927  
 <211> 30  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 heavy chain variable framework 1 (FW1) sequence

<400> 927  
 Glu Val Glu Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly  
 1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Thr Ser Gly Phe Thr Phe Thr  
 20 25 30

<210> 928  
 <211> 15  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 928  
 gatcactaca tgagc 15

<210> 929  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 929  
 Asp His Tyr Met Ser  
 1 5

<210> 930  
 <211> 42  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 heavy chain variable framework 2 (FW2) sequence  
  
 <400> 930  
 tgggtccgcc agcctccagg aaaggcactt gagtggttgg ga 42

<210> 931  
 <211> 14  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 heavy chain variable framework 2 (FW2) sequence  
  
 <400> 931  
  
 Trp Val Arg Gln Pro Pro Gly Lys Ala Leu Glu Trp Leu Gly  
 1 5 10

<210> 932  
 <211> 57  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 932  
 tttattagaa acaaagctaa tggttacaca acagagtaca gtgcatctgt gaagggt 57

<210> 933  
 <211> 19  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 933  
  
 Phe Ile Arg Asn Lys Ala Asn Gly Tyr Thr Thr Glu Tyr Ser Ala Ser  
 1 5 10 15  
  
 Val Lys Gly

<210> 934  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 heavy chain variable framework 3 (FW3) sequence  
  
 <400> 934  
 cggttcacca tctccagaga taattcccaa agcatcctct atcttcaaat gaaaaccctg 60  
 agaactgagg acagtgccac ttattactgt gcaaga 96

<210> 935  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 heavy chain variable framework 3 (FW3)  
sequence

<400> 935

Arg Phe Thr Ile Ser Arg Asp Asn Ser Gln Ser Ile Leu Tyr Leu Gln  
1 5 10 15

Met Lys Thr Leu Arg Thr Glu Asp Ser Ala Thr Tyr Tyr Cys Ala Arg  
20 25 30

<210> 936  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 936  
ccttctgact gggactcctg gtttgcttac 30

<210> 937  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 937

Pro Ser Asp Trp Asp Ser Trp Phe Ala Tyr  
1 5 10

<210> 938  
<211> 33  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 heavy chain variable framework 4 (FW4)  
sequence

<400> 938  
tggggccaag ggactctggt cactgtctct gca 33

<210> 939  
<211> 11  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 heavy chain variable framework 4 (FW4)



# sequence

<400> 939

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
1 5 10

<210> 940

<211> 393

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 8A9 Light chain: DNA sequence (393 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 940

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atgaagttgc ctgttaggct gttggtgctg atgttctgga ttcctgcttc cagcagtgat      60
gttttgatga cccaaactcc actctccctg cctgtcagtc ttggtgatca agcctccatc      120
tcttcgagat ctagtcagag cattgtacat agtaatggca acacctatct agattggtac      180
ttgcagaaac caggccagtc tccaaagctc ctgatctaca gagtttccaa ccgattttct      240
gggggtcccag acaggttcag tggcagtgga tcaggacag atttcacact caagatcagc      300
agagtgaggg ctgaggatct gggactttat tactgttttc aagggttcaca tgttccgtgg      360
gcgttcggtg gaggcaccaa gctggaaatc aaa                                  393
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<210> 941

<211> 131

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 8A9 Light chain: Amino acid sequence (131 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 941

Met Lys Leu Pro Val Arg Leu Leu Val Leu Met Phe Trp Ile Pro Ala  
1 5 10 15

Ser Ser Ser Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val  
20 25 30

Ser Leu Gly Asp Gln Ala Ser Ile Ser Cys Arg Ser Ser Gln Ser Ile  
35 40 45

Val His Ser Asn Gly Asn Thr Tyr Leu Asp Trp Tyr Leu Gln Lys Pro  
50 55 60

Gly Gln Ser Pro Lys Leu Leu Ile Tyr Arg Val Ser Asn Arg Phe Ser  
65 70 75 80

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
85 90 95

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Leu Tyr Tyr Cys

100

105

110

Phe Gln Gly Ser His Val Pro Trp Ala Phe Gly Gly Gly Thr Lys Leu  
 115 120 125

Glu Ile Lys  
 130

<210> 942  
 <211> 69  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 light chain variable framework 1 (FW1)  
 sequence

<400> 942  
 gatgttttga tgacccaaac tccactctcc ctgcctgtca gtcttggtga tcaagcctcc 60  
 atctcttgc 69

<210> 943  
 <211> 23  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 light chain variable framework 1 (FW1)  
 sequence

<400> 943

Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
 1 5 10 15

Asp Gln Ala Ser Ile Ser Cys  
 20

<210> 944  
 <211> 48  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 light chain variable complementarity  
 determining regions 1 (CDR1) sequence

<400> 944  
 agatctagtc agagcattgt acatagtaat ggcaacacct atttagat 48

<210> 945  
 <211> 16  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 light chain variable complementarity  
 determining regions 1 (CDR1) sequence

<400> 945

Arg Ser Ser Gln Ser Ile Val His Ser Asn Gly Asn Thr Tyr Leu Asp  
1 5 10 15

<210> 946  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 light chain variable framework 2 (FW2)  
sequence

<400> 946  
tgggtacttgc agaaaccagg ccagtctcca aagctcctga tctac 45

<210> 947  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 light chain variable framework 2 (FW2)  
sequence

<400> 947

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 948  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 948  
agagtttcca accgattttc t 21

<210> 949  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 949

Arg Val Ser Asn Arg Phe Ser  
1 5

<210> 950  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 8A9 light chain variable framework 3 (FW3)  
sequence

<400> 950  
 ggggtcccag acaggttcag tggcagtgga tcagggacag atttcacact caagatcagc 60  
 agagtggagg ctgaggatct gggactttat tactgt 96

<210> 951  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 light chain variable framework 3 (FW3) sequence

<400> 951  
 Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 1 5 10 15  
 Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Leu Tyr Tyr Cys  
 20 25 30

<210> 952  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 952  
 tttcaagggtt cacatgttcc gtgggcg 27

<210> 953  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 953  
 Phe Gln Gly Ser His Val Pro Trp Ala  
 1 5

<210> 954  
 <211> 30  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 8A9 light chain variable framework 4 (FW4) sequence

<400> 954  
 ttcggtggag gcaccaagct ggaaatcaaa 30

<210> 955  
 <211> 10  
 <212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 8A9 light chain variable framework 4 (FW4) sequence

<400> 955

Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
1 5 10

<210> 956

<211> 399

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 18G12 Heavy chain: DNA sequence (399 bp) - Signal sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 956

atgggatgga gctatatcat cctctttttg gtcgcaacag ctacaggtgt ccactcccag 60  
gtccaactgc agcagtctgg ggctgaactg gtgaagcctg gggcttcagt gaagttgtcc 120  
tgcaaggcct ctggctacac cttcaccggc tactttttgt actgggtgaa gcagaggcct 180  
ggacaaggcc ttgagtggat tggggggatt aatcctgaca atggtggtat tgacttcaat 240  
gagaagttca ggaacaaggc cacactgact gtagacaaat cctccagcac agcctacatg 300  
caactcagca gcctgacatc tgaggactct gcggtctatt attgtacatt actaataggg 360  
aactattggg gccaaggcac cactctcaca gtctcctca 399

<210> 957

<211> 133

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 18G12 Heavy chain: Amino acid sequence (133 aa) - Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 957

Met Gly Trp Ser Tyr Ile Ile Leu Phe Leu Val Ala Thr Ala Thr Gly  
1 5 10 15

Val His Ser Gln Val Gln Leu Gln Gln Ser Gly Ala Glu Leu Val Lys  
20 25 30

Pro Gly Ala Ser Val Lys Leu Ser Cys Lys Ala Ser Gly Tyr Thr Phe  
35 40 45

Thr Gly Tyr Phe Leu Tyr Trp Val Lys Gln Arg Pro Gly Gln Gly Leu  
50 55 60

Glu Trp Ile Gly Gly Ile Asn Pro Asp Asn Gly Gly Ile Asp Phe Asn  
65 70 75 80

Glu Lys Phe Arg Asn Lys Ala Thr Leu Thr Val Asp Lys Ser Ser Ser

85

90

95

Thr Ala Tyr Met Gln Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val  
 100 105 110

Tyr Tyr Cys Thr Leu Leu Ile Gly Asn Tyr Trp Gly Gln Gly Thr Thr  
 115 120 125

Leu Thr Val Ser Ser  
 130

<210> 958

<211> 90

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 18G12 heavy chain variable framework 1 (FW1) sequence

<400> 958

caggccaac tgcagcagtc tggggctgaa ctggtgaagc ctggggcttc agtgaagttg 60

tcctgcaagg cttctggcta caccttcacc 90

<210> 959

<211> 30

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 18G12 heavy chain variable framework 1 (FW1) sequence

<400> 959

Gln Val Gln Leu Gln Gln Ser Gly Ala Glu Leu Val Lys Pro Gly Ala  
 1 5 10 15

Ser Val Lys Leu Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr  
 20 25 30

<210> 960

<211> 15

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 960

ggctactttt tgtac 15

<210> 961

<211> 5

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 39H5 heavy chain variable complementarity

determining regions 1 (CDR1) sequence

<400> 961

Gly Tyr Phe Leu Tyr  
1 5

<210> 962

<211> 42

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 18G12 heavy chain variable framework 2 (FW2)  
sequence

<400> 962

tgggtgaagc agaggcctgg acaaggcctt gattggattg gg

42

<210> 963

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 18G12 heavy chain variable framework 2 (FW2)  
sequence

<400> 963

Trp Val Lys Gln Arg Pro Gly Gln Gly Leu Glu Trp Ile Gly  
1 5 10

<210> 964

<211> 51

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 18G12 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 964

gggattaatc ctgacaatgg tggattgac ttcaatgaga agttcaggaa c

51

<210> 965

<211> 17

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 18G12 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 965

Gly Ile Asn Pro Asp Asn Gly Gly Ile Asp Phe Asn Glu Lys Phe Arg  
1 5 10 15

Asn

<210> 966

<211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 18G12 heavy chain variable framework 3 (FW3) sequence  
  
 <400> 966  
 aaggccacac tgactgtaga caaatcctcc agcacagcct acatgcaact cagcagcctg 60  
  
 acatctgagg actctgcggt ctattattgt acatta 96  
  
 <210> 967  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 18G12 heavy chain variable framework 3 (FW3) sequence  
  
 <400> 967  
  
 Lys Ala Thr Leu Thr Val Asp Lys Ser Ser Ser Thr Ala Tyr Met Gln  
 1 5 10 15  
  
 Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val Tyr Tyr Cys Thr Leu  
 20 25 30  
  
 <210> 968  
 <211> 15  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 18G12 heavy chain variable complementarity determining regions 3 (CDR3) sequence  
  
 <400> 968  
 ctaatagga actat 15  
  
 <210> 969  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 18G12 heavy chain variable complementarity determining regions 3 (CDR3) sequence  
  
 <400> 969  
  
 Leu Ile Gly Asn Tyr  
 1 5  
  
 <210> 970  
 <211> 33  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 18G12 heavy chain variable framework 4 (FW4) sequence



<400> 970  
tggggccaag gcaccactct cacagtctcc tca 33

<210> 971  
<211> 11  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 heavy chain variable framework 4 (FW4)  
sequence

<400> 971  
Trp Gly Gln Gly Thr Thr Leu Thr Val Ser Ser  
1 5 10

<210> 972  
<211> 393  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 Light chain: DNA sequence (393 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 972  
atgagtcctg cccagttcct gtttctgtta gtgctctgga ttcgggaaac caatggtgat 60  
gttgtgatga cccagactcc actcactttg tcggtaacca ttggacagcc agcctccatc 120  
tcttgcaagt caagtcagag cctcttacat agtgatggaa agacatattt gatttggttg 180  
ttacagaggc caggccagtc tccaaagcgc ctaatctatc tgggtgtctaa actggactct 240  
ggagtccctg acaggtttcac tggcagtgga tcagggacag atttcacact gaaaatcagc 300  
agagtggagg ctgaggattt gggagtttat ttttgctgtc aaggtaaca ttttccgtgg 360  
acgttcggtg gaggcacat gctggaaatc aaa 393

<210> 973  
<211> 131  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 Light chain: Amino acid sequence (131 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 973  
Met Ser Pro Ala Gln Phe Leu Phe Leu Val Leu Trp Ile Arg Glu  
1 5 10 15  
Thr Asn Gly Asp Val Val Met Thr Gln Thr Pro Leu Thr Leu Ser Val  
20 25 30  
Thr Ile Gly Gln Pro Ala Ser Ile Ser Cys Lys Ser Ser Gln Ser Leu  
35 40 45  
Leu His Ser Asp Gly Lys Thr Tyr Leu Ile Trp Leu Leu Gln Arg Pro

50

55

60

Gly Gln Ser Pro Lys Arg Leu Ile Tyr Leu Val Ser Lys Leu Asp Ser  
65 70 75 80

Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr  
85 90 95

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys  
100 105 110

Cys Gln Gly Thr His Phe Pro Trp Thr Phe Gly Gly Gly Thr Met Leu  
115 120 125

Glu Ile Lys  
130

<210> 974  
<211> 69  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable framework 1 (FW1)  
sequence

<400> 974  
gatgttgtga tgaccagac tccactcact ttgtcggtaa ccattggaca gccagcctcc 60  
atctcttgc 69

<210> 975  
<211> 23  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable framework 1 (FW1)  
sequence

<400> 975

Asp Val Val Met Thr Gln Thr Pro Leu Thr Leu Ser Val Thr Ile Gly  
1 5 10 15

Gln Pro Ala Ser Ile Ser Cys  
20

<210> 976  
<211> 48  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 976  
aagtaagtc agagcctctt acatagtgat ggaaagacat atttgatt 48

<210> 977  
<211> 16  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 977

Lys Ser Ser Gln Ser Leu Leu His Ser Asp Gly Lys Thr Tyr Leu Ile  
1 5 10 15

<210> 978  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable framework 2 (FW2)  
sequence

<400> 978  
tggttggttac agaggccagg ccagtctcca aagcgctaa tctat 45

<210> 979  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable framework 2 (FW2)  
sequence

<400> 979

Trp Leu Leu Gln Arg Pro Gly Gln Ser Pro Lys Arg Leu Ile Tyr  
1 5 10 15

<210> 980  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 980  
ctggtgtcta aactggactc t 21

<210> 981  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 981

Leu Val Ser Lys Leu Asp Ser

1 5

<210> 982  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable framework 3 (FW3)  
sequence

<400> 982  
ggagtccctg acaggttcac tggcagtgga tcagggacag atttcacact gaaaatcagc 60  
agagtggagg ctgaggattt gggagtttat ttttgc 96

<210> 983  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable framework 3 (FW3)  
sequence

<400> 983

Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys  
20 25 30

<210> 984  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 984  
tgtcaaggta cacattttcc gtggacg 27

<210> 985  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18G12 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 985

Cys Gln Gly Thr His Phe Pro Trp Thr  
1 5

<210> 986  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
 <223> Mouse antibody 18G12 light chain variable framework 4 (FW4) sequence

<400> 986  
 ttcggtggag gcaccatgct ggaaatcaaa 30

<210> 987  
 <211> 10  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 18G12 light chain variable framework 4 (FW4) sequence

<400> 987

Phe Gly Gly Gly Thr Met Leu Glu Ile Lys  
 1 5 10

<210> 988  
 <211> 416  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 20A10 Heavy chain: DNA sequence (417 bp) - Signal sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 988  
 atgaacttcg ggttcagctt gatcttcctt gtccttggtt taaaagggtg ccagtgtgaa 60  
 gtgatgctgg tggagtctgg gggaggctta gtgaagcctg gagggtcctt gaaactctcc 120  
 tgtcagcctc tggattcact ttcagtacct atgccatgtc ttggattcgc cagactccag 180  
 agaagaggct ggagtgggtc gcatccattg gtcgtgctgg ttccacctac tattcagaca 240  
 gtgtgaaggg ccgattcacc atctccagag ataatgtccg gaacatcctg tacctgcaaa 300  
 tgagcagtct gaggtctgag gacacggcca tgtattactg tgctagaggc ccgatctaca 360  
 atgattacga cgagtttgct tactggggcc aagggaactct ggtcactgtc tctgca 416

<210> 989  
 <211> 139  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 20A10 Heavy chain: Amino acid sequence (139 aa) - Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 989

Met Asn Phe Gly Phe Ser Leu Ile Phe Leu Val Leu Val Leu Lys Gly  
 1 5 10 15

Val Gln Cys Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu Val Lys  
 20 25 30

Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe

35

40

45

Ser Thr Tyr Ala Met Ser Trp Ile Arg Gln Thr Pro Glu Lys Arg Leu  
 50 55 60

Glu Trp Val Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr Ser Asp  
 65 70 75 80

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Val Arg Asn Ile  
 85 90 95

Leu Tyr Leu Gln Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Met Tyr  
 100 105 110

Tyr Cys Ala Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe Ala Tyr  
 115 120 125

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
 130 135

&lt;210&gt; 990

&lt;211&gt; 90

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Mouse antibody 20A10 heavy chain variable framework 1 (FW1) sequence

&lt;400&gt; 990

gaagtgatgc tgggtggagtc tgggggaggc ttagtgaagc ctggagggtc cctgaaactc 60

tcctgtgcag cctctggatt cactttcagt 90

&lt;210&gt; 991

&lt;211&gt; 30

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Mouse antibody 20A10 heavy chain variable framework 1 (FW1) sequence

&lt;400&gt; 991

Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
 20 25 30

&lt;210&gt; 992

&lt;211&gt; 15

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Mouse antibody 20A10 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 992  
acctatgccca tgtct

15

<210> 993  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 993

Thr Tyr Ala Met Ser  
1 5

<210> 994  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 heavy chain variable framework 2 (FW2)  
sequence

<400> 994  
tggattcgcc agactccaga gaagaggctg gagtgggtcg ca

42

<210> 995  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 heavy chain variable framework 2 (FW2)  
sequence

<400> 995

Trp Ile Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val Ala  
1 5 10

<210> 996  
<211> 48  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 996  
tccattggtc gtgctggttc cacctactat tcagacagtg tgaagggc

48

<210> 997  
<211> 16  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 997

Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr Ser Asp Ser Val Lys Gly  
1 5 10 15

<210> 998

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 20A10 heavy chain variable framework 3 (FW3)  
sequence

<400> 998

cgattcacca tctccagaga taatgtccgg aacatcctgt acctgcaaat gagcagtcgt 60

aggctctgagg acacggccat gtattactgt gctaga 96

<210> 999

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 20A10 heavy chain variable framework 3 (FW3)  
sequence

<400> 999

Arg Phe Thr Ile Ser Arg Asp Asn Val Arg Asn Ile Leu Tyr Leu Gln  
1 5 10 15

Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Met Tyr Tyr Cys Ala Arg  
20 25 30

<210> 1000

<211> 36

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 20A10 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1000

ggccccgatct acaatgatta cgacgagttt gcttac 36

<210> 1001

<211> 12

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 20A10 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1001

Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe Ala Tyr  
1 5 10



<210> 1002  
 <211> 33  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 20A10 heavy chain variable framework 4 (FW4) sequence  
  
 <400> 1002  
 tggggccaag ggactctggt cactgtctct gca 33

<210> 1003  
 <211> 11  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 20A10 heavy chain variable framework 4 (FW4) sequence

<400> 1003  
  
 Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
 1 5 10

<210> 1004  
 <211> 396  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 20A10 Light chain: DNA sequence (396 bp) - Signal sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1004  
 atggaatcac agactcaggt cttcctctcc ctgctgctct gggatatctgg tacctgtggg 60  
  
 aacattatga tgacacagtc gccatcatct ctggctgtgt ctgcaggaga aaaggtcact 120  
  
 atgagctgta agtccagtca aagtgtttta tacagttcaa atcagaagaa ctatttggcc 180  
  
 tggtagcagc agaaaccagg gcagtctcct aaactgctga tctactgggc atccactagg 240  
  
 gaatctggtg tccctgatcg cttcacaggc agtggatctg ggacagattt tactcttacc 300  
  
 atcagcagtg tacaagctga agacctggca gtttattact gtcataata cctctcctcg 360  
  
 ctcacgttcg gtgctgggac caagctggag ctgaaa 396

<210> 1005  
 <211> 132  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 20A10 Light chain: Amino acid sequence (132 aa) - Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1005  
  
 Met Glu Ser Gln Thr Gln Val Phe Leu Ser Leu Leu Trp Val Ser  
 1 5 10 15

Gly Thr Cys Gly Asn Ile Met Met Thr Gln Ser Pro Ser Ser Leu Ala

20

25

30

Val Ser Ala Gly Glu Lys Val Thr Met Ser Cys Lys Ser Ser Gln Ser  
 35 40 45

Val Leu Tyr Ser Ser Asn Gln Lys Asn Tyr Leu Ala Trp Tyr Gln Gln  
 50 55 60

Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr Trp Ala Ser Thr Arg  
 65 70 75 80

Glu Ser Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser Gly Thr Asp  
 85 90 95

Phe Thr Leu Thr Ile Ser Ser Val Gln Ala Glu Asp Leu Ala Val Tyr  
 100 105 110

Tyr Cys His Gln Tyr Leu Ser Ser Leu Thr Phe Gly Ala Gly Thr Lys  
 115 120 125

Leu Glu Leu Lys  
 130

<210> 1006

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 20A10 light chain variable framework 1 (FW1)  
 sequence

<400> 1006

aacattatga tgacacagtc gccatcatct ctggctgtgt ctgcaggaga aaaggtcact 60

atgagctgt 69

<210> 1007

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 20A10 light chain variable framework 1 (FW1)  
 sequence

<400> 1007

Asn Ile Met Met Thr Gln Ser Pro Ser Ser Leu Ala Val Ser Ala Gly  
 1 5 10 15

Glu Lys Val Thr Met Ser Cys  
 20

<210> 1008

<211> 51

<212> DNA

<213> Artificial Sequence

<220>  
 <223> Mouse antibody 20A10 light chain variable complementarity  
 determining regions 1 (CDR1) sequence

<400> 1008  
 aagtcacgctc aaagtgtttt atacagttca aatcagaaga actatttggc c 51

<210> 1009  
 <211> 17  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 20A10 light chain variable complementarity  
 determining regions 1 (CDR1) sequence

<400> 1009

Lys	Ser	Ser	Gln	Ser	Val	Leu	Tyr	Ser	Ser	Asn	Gln	Lys	Asn	Tyr	Leu
1				5					10					15	

Ala

<210> 1010  
 <211> 45  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 20A10 light chain variable framework 2 (FW2)  
 sequence

<400> 1010  
 tgggtaccagc agaaaccagg gcagtctcct aaactgctga tctac 45

<210> 1011  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 20A10 light chain variable framework 2 (FW2)  
 sequence

<400> 1011

Trp	Tyr	Gln	Gln	Lys	Pro	Gly	Gln	Ser	Pro	Lys	Leu	Leu	Ile	Tyr
1				5					10					15

<210> 1012  
 <211> 21  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 20A10 light chain variable complementarity  
 determining regions 2 (CDR2) sequence

<400> 1012  
 tgggcatcca ctagggaatc t 21

<210> 1013  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1013

Trp Ala Ser Thr Arg Glu Ser  
1 5

<210> 1014  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 light chain variable framework 3 (FW3)  
sequence

<400> 1014  
gggtgccctg atcgcttcac aggcagtgga tctgggacag attttactct taccatcagc 60  
agtgtacaag ctgaagacct ggcagtttat tactgt 96

<210> 1015  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 light chain variable framework 3 (FW3)  
sequence

<400> 1015

Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Thr Ile Ser Ser Val Gln Ala Glu Asp Leu Ala Val Tyr Tyr Cys  
20 25 30

<210> 1016  
<211> 24  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1016  
catcaatacc tctcctcgct cacg 24

<210> 1017  
<211> 8  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 20A10 light chain variable complementarity

determining regions 3 (CDR3) sequence

<400> 1017

His Gln Tyr Leu Ser Ser Leu Thr  
1 5

<210> 1018

<211> 30

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 20A10 light chain variable framework 4 (FW4)  
sequence

<400> 1018

ttcgggtgctg ggaccaagct ggagctgaaa 30

<210> 1019

<211> 10

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 20A10 light chain variable framework 4 (FW4)  
sequence

<400> 1019

Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys  
1 5 10

<210> 1020

<211> 413

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 25E6 Heavy chain: DNA sequence (414 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1020

atgaacttcg ggctcagctt gattttcctt gccctcattt taaaagggtg ccagtgtgag 60

gtgcagctgg tggagtctgg gggagactta gtgaagcctg gagggtcctt gaaactctcc 120

tgtgcagcct ctggtttcac ttctagtagt tatggaatgt cttgggttcg ccagactcca 180

gacaagaggc tggagtgggt cgcaaccatt agtaatgggt gtagacacac cttctatcca 240

gacagtgtga aggggcgatt caccatctcc agagacaatg ccaagaacac cctgtatctg 300

caaatgagca gtctgaagtt gaggacacag ccatgtatct atgtgtaaga cagactggga 360

cggagggtct gtttgcttac tggggccaag ggactctggt cactgtctct gca 413

<210> 1021

<211> 138

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 25E6 Heavy chain: Amino acid sequence (138 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1021

Met Asn Phe Gly Leu Ser Leu Ile Phe Leu Ala Leu Ile Leu Lys Gly  
1 5 10 15

Val Gln Cys Glu Val Gln Leu Val Glu Ser Gly Gly Asp Leu Val Lys  
20 25 30

Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe  
35 40 45

Ser Ser Tyr Gly Met Ser Trp Val Arg Gln Thr Pro Asp Lys Arg Leu  
50 55 60

Glu Trp Val Ala Thr Ile Ser Asn Gly Gly Arg His Thr Phe Tyr Pro  
65 70 75 80

Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn  
85 90 95

Thr Leu Tyr Leu Gln Met Ser Ser Leu Lys Ser Glu Asp Thr Ala Met  
100 105 110

Tyr Leu Cys Val Arg Gln Thr Gly Thr Glu Gly Trp Phe Ala Tyr Trp  
115 120 125

Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
130 135

<210> 1022

<211> 90

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 25E6 heavy chain variable framework 1 (FW1)  
sequence

<400> 1022

gaggtgcagc tgggtggagtc tgggggagac ttagtgaagc ctggagggtc cctgaaactc 60

tcctgtgcag cctctggttt cactttcagt 90

<210> 1023

<211> 30

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 25E6 heavy chain variable framework 1 (FW1)  
sequence

<400> 1023

Glu Val Gln Leu Val Glu Ser Gly Gly Asp Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 1024  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1024  
agttatggaa tgtct 15

<210> 1025  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1025

Ser Tyr Gly Met Ser  
1 5

<210> 1026  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 heavy chain variable framework 2 (FW2)  
sequence

<400> 1026  
tgggttcgcc agactccaga caagaggctg gagtgggtcg ca 42

<210> 1027  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 heavy chain variable framework 2 (FW2)  
sequence

<400> 1027

Trp Val Arg Gln Thr Pro Asp Lys Arg Leu Glu Trp Val Ala  
1 5 10

<210> 1028  
<211> 51  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1028  
accattagta atggtggttag acacaccttc tatccagaca gtgtgaaggg g 51

<210> 1029  
<211> 17  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1029  
Thr Ile Ser Asn Gly Gly Arg His Thr Phe Tyr Pro Asp Ser Val Lys  
1 5 10 15

Gly

<210> 1030  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 heavy chain variable framework 3 (FW3)  
sequence

<400> 1030  
cgattcacca tctccagaga caatgccaaag aacaccctgt atctgcaaat gagcagtcctg 60  
aagtctgagg acacagccat gtatttatgt gtaaga 96

<210> 1031  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 heavy chain variable framework 3 (FW3)  
sequence

<400> 1031  
Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln  
1 5 10 15

Met Ser Ser Leu Lys Ser Glu Asp Thr Ala Met Tyr Leu Cys Val Arg  
20 25 30

<210> 1032  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1032  
cagactggga cggagggctg gtttgcttac 30



<210> 1033  
 <211> 10  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 25E6 heavy chain variable complementarity  
 determining regions 3 (CDR3) sequence

<400> 1033

Gln Thr Gly Thr Glu Gly Trp Phe Ala Tyr  
 1 5 10

<210> 1034  
 <211> 33  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 25E6 heavy chain variable framework 4 (FW4)  
 sequence

<400> 1034

tggggccaag ggactctggt cactgtctct gca 33

<210> 1035  
 <211> 11  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 25E6 heavy chain variable framework 4 (FW4)  
 sequence

<400> 1035

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
 1 5 10

<210> 1036  
 <211> 392  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 25E6 Light chain: DNA sequence (393 bp) - Signal  
 sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1036

atgagtcctg cccagttcct gtttctgtta gtgctctgga ttcgggaaac caacggtgat 60  
  
 gttgtgatga cccagactcc actcactttg tcggttacca ttggacaacc agcctccatc 120  
  
 tcttgcaagt caagtcagag cctcttagat agtgatggaa agacatattt gaattggttg 180  
  
 ttacagaggc caggccagtc tccaaagcgc ctaatctatc tgggtgtctaa actggactct 240  
  
 ggagtccttg acaggttcac tggcagtgga tcagggacag atttcacact gaaaatcagc 300  
  
 agagggaggc tgaggatttg ggagtttatt attgctggca aggtacacat tttcctcaga 360  
  
 cgttcggtgg aggaccaag ctggaaatca aa 392

<210> 1037  
 <211> 131  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 25E6 Light chain: Amino acid sequence (131 aa) -  
 Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1037

Met Ser Pro Ala Gln Phe Leu Phe Leu Leu Val Leu Trp Ile Arg Glu  
 1 5 10 15

Thr Asn Gly Asp Val Val Met Thr Gln Thr Pro Leu Thr Leu Ser Val  
 20 25 30

Thr Ile Gly Gln Pro Ala Ser Ile Ser Cys Lys Ser Ser Gln Ser Leu  
 35 40 45

Leu Asp Ser Asp Gly Lys Thr Tyr Leu Asn Trp Leu Leu Gln Arg Pro  
 50 55 60

Gly Gln Ser Pro Lys Arg Leu Ile Tyr Leu Val Ser Lys Leu Asp Ser  
 65 70 75 80

Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 85 90 95

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
 100 105 110

Trp Gln Gly Thr His Phe Pro Gln Thr Phe Gly Gly Gly Thr Lys Leu  
 115 120 125

Glu Ile Lys  
 130

<210> 1038  
 <211> 69  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 25E6 light chain variable framework 1 (FW1)  
 sequence

<400> 1038  
 gatgttgatga tgacccagac tccactcact ttgtcgggta ccattggaca accagcctcc 60  
 atctcttgcc 69

<210> 1039  
 <211> 23  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 25E6 light chain variable framework 1 (FW1)

sequence

<400> 1039

Asp Val Val Met Thr Gln Thr Pro Leu Thr Leu Ser Val Thr Ile Gly  
1 5 10 15

Gln Pro Ala Ser Ile Ser Cys  
20

<210> 1040

<211> 48

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 25E6 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1040

aagtcaagtc agagcctctt agatagtgat ggaaagacat atttgaat 48

<210> 1041

<211> 16

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 25E6 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1041

Lys Ser Ser Gln Ser Leu Leu Asp Ser Asp Gly Lys Thr Tyr Leu Asn  
1 5 10 15

<210> 1042

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 25E6 light chain variable framework 2 (FW2)  
sequence

<400> 1042

tgggtgttac agaggccagg ccagtctcca aagcgcctaa tctat 45

<210> 1043

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 25E6 light chain variable framework 2 (FW2)  
sequence

<400> 1043

Trp Leu Leu Gln Arg Pro Gly Gln Ser Pro Lys Arg Leu Ile Tyr  
1 5 10 15

<210> 1044

<211> 21  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 25E6 light chain variable complementarity  
 determining regions 2 (CDR2) sequence  
  
 <400> 1044  
 ctggtgtcta aactggactc t 21

<210> 1045  
 <211> 7  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 25E6 light chain variable complementarity  
 determining regions 2 (CDR2) sequence  
  
 <400> 1045  
  
 Leu Val Ser Lys Leu Asp Ser  
 1 5

<210> 1046  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 25E6 light chain variable framework 3 (FW3)  
 sequence  
  
 <400> 1046  
 ggagtcctg acaggttcac tggcagtgga tcaggacag atttcacact gaaaatcagc 60  
 agagtggagg ctgaggattt gggagtttat tattgc 96

<210> 1047  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 25E6 light chain variable framework 3 (FW3)  
 sequence  
  
 <400> 1047  
  
 Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
 20 25 30

<210> 1048  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 25E6 light chain variable complementarity  
 determining regions 3 (CDR3) sequence

<400> 1048  
tggcaaggta cacattttcc tcagacg 27

<210> 1049  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1049  
Trp Gln Gly Thr His Phe Pro Gln Thr  
1 5

<210> 1050  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 light chain variable framework 4 (FW4)  
sequence

<400> 1050  
ttcgggtggag gcaccaagct ggaaatcaaa 30

<210> 1051  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 25E6 light chain variable framework 4 (FW4)  
sequence

<400> 1051  
Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
1 5 10

<210> 1052  
<211> 399  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 Heavy chain: DNA sequence (399 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1052  
atgggatgga gctatatcat cctctttttg gtagcaacag ctacaggtgt ccaactcccag 60  
gtccaactgc agcagcctgg ggctgaactg gtgcagcctg gggcttcagt gaagttgtcc 120  
tgcaaggctt ctggctacac cttcaccggc tactttttgt actgggtgaa gcagaggcct 180  
ggacatggcc ttgagtggat tgggggaatt catcctagca atgggtgatac tgacttcaat 240  
gagaagtcca agaacaaggc cacactgact gtagacatat cctccagcac tgcctacatg 300  
caactcagca gcctgacatc tgaggactct gcggtctatt attgtacatt actaataggg 360

gtctactggg gccaaaggcac cactctcaca gtctcctca

399

<210> 1053  
<211> 133  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 Heavy chain: Amino acid sequence (133 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1053

Met Gly Trp Ser Tyr Ile Ile Leu Phe Leu Val Ala Thr Ala Thr Gly  
1 5 10 15

Val His Ser Gln Val Gln Leu Gln Gln Pro Gly Ala Glu Leu Val Gln  
20 25 30

Pro Gly Ala Ser Val Lys Leu Ser Cys Lys Ala Ser Gly Tyr Thr Phe  
35 40 45

Thr Gly Tyr Phe Leu Tyr Trp Val Lys Gln Arg Pro Gly His Gly Leu  
50 55 60

Glu Trp Ile Gly Gly Ile His Pro Ser Asn Gly Asp Thr Asp Phe Asn  
65 70 75 80

Glu Lys Phe Lys Asn Lys Ala Thr Leu Thr Val Asp Ile Ser Ser Ser  
85 90 95

Thr Ala Tyr Met Gln Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val  
100 105 110

Tyr Tyr Cys Thr Leu Leu Ile Gly Val Tyr Trp Gly Gln Gly Thr Thr  
115 120 125

Leu Thr Val Ser Ser  
130

<210> 1054  
<211> 90  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 heavy chain variable framework 1 (FW1)  
sequence

<400> 1054

caggtccaac tgcagcagcc tggggctgaa ctggtgcagc ctggggcttc agtgaagttg 60

tcctgcaagg cttctggcta caccttcacc 90

<210> 1055  
<211> 30  
<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 28F9 heavy chain variable framework 1 (FW1) sequence

<400> 1055

Gln Val Gln Leu Gln Gln Pro Gly Ala Glu Leu Val Gln Pro Gly Ala  
1 5 10 15

Ser Val Lys Leu Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr  
20 25 30

<210> 1056

<211> 15

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 28F9 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1056

ggctactttt tgtac

15

<210> 1057

<211> 5

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 28F9 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1057

Gly Tyr Phe Leu Tyr  
1 5

<210> 1058

<211> 42

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 28F9 heavy chain variable framework 2 (FW2) sequence

<400> 1058

tgggtgaagc agaggcctgg acatggcctt gaggaggattg gg

42

<210> 1059

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 28F9 heavy chain variable framework 2 (FW2) sequence

<400> 1059

Trp Val Lys Gln Arg Pro Gly His Gly Leu Glu Trp Ile Gly

1 5 10

<210> 1060  
<211> 51  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1060  
ggaattcatc ctagcaatgg tgatactgac ttcaatgaga agttcaagaa c 51

<210> 1061  
<211> 17  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1061

Gly Ile His Pro Ser Asn Gly Asp Thr Asp Phe Asn Glu Lys Phe Lys  
1 5 10 15

Asn

<210> 1062  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 heavy chain variable framework 3 (FW3)  
sequence

<400> 1062  
aaggccacac tgactgtaga catatcctcc agcactgcct acatgcaact cagcagcctg 60

acatctgagg actctgcggt ctattattgt acatta 96

<210> 1063  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 heavy chain variable framework 3 (FW3)  
sequence

<400> 1063

Lys Ala Thr Leu Thr Val Asp Ile Ser Ser Ser Thr Ala Tyr Met Gln  
1 5 10 15

Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val Tyr Tyr Cys Thr Leu  
20 25 30



<210> 1064  
 <211> 15  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 28F9 heavy chain variable complementarity  
 determining regions 3 (CDR3) sequence  
  
 <400> 1064  
 ctaatagggg tctac 15  
  
 <210> 1065  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 28F9 heavy chain variable complementarity  
 determining regions 3 (CDR3) sequence  
  
 <400> 1065  
  
 Leu Ile Gly Val Tyr  
 1 5  
  
 <210> 1066  
 <211> 33  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 28F9 heavy chain variable framework 4 (FW4)  
 sequence  
  
 <400> 1066  
 tggggccaag gcaccactct cacagtctcc tca 33  
  
 <210> 1067  
 <211> 11  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 28F9 heavy chain variable framework 4 (FW4)  
 sequence  
  
 <400> 1067  
  
 Trp Gly Gln Gly Thr Thr Leu Thr Val Ser Ser  
 1 5 10  
  
 <210> 1068  
 <211> 393  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 28F9 Light chain: DNA sequence (393 bp) - Signal  
 sequenceFR1CDR1FR2CDR2FR3CDR3FR4  
  
 <400> 1068  
 atgagtcctg cccagttcct gtttctgtta gtgctctgga ttcgggaaac caacggtgat 60  
 gttgtgatga cccagactcc actcactttg tcggttacca ttggacaacc agcctccatc 120

tcttgcaagt caagtcagag cctcttacat agtgatggaa agacatattt gatttggttg	180
ttacagaggc caggccagtc tccaaagcgc ctaatctatc tggtgtctaa actggactct	240
ggagtccctg acaggttcac cggcagtgga tcagggacag atttcacact gaaaatcagc	300
agagtggagg ctgaggattt gggagtttat ttttgctgtc aaggtacaca ttttccgtgg	360
acgttcggtg gaggcaccat gctggaaatc aaa	393

<210> 1069  
 <211> 131  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 28F9 Light chain: Amino acid sequence (131 aa) -  
 Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1069

Met	Ser	Pro	Ala	Gln	Phe	Leu	Phe	Leu	Leu	Val	Leu	Trp	Ile	Arg	Glu
1				5					10					15	

Thr	Asn	Gly	Asp	Val	Val	Met	Thr	Gln	Thr	Pro	Leu	Thr	Leu	Ser	Val
			20					25					30		

Thr	Ile	Gly	Gln	Pro	Ala	Ser	Ile	Ser	Cys	Lys	Ser	Ser	Gln	Ser	Leu
		35					40					45			

Leu	His	Ser	Asp	Gly	Lys	Thr	Tyr	Leu	Ile	Trp	Leu	Leu	Gln	Arg	Pro
	50					55					60				

Gly	Gln	Ser	Pro	Lys	Arg	Leu	Ile	Tyr	Leu	Val	Ser	Lys	Leu	Asp	Ser
65					70					75				80	

Gly	Val	Pro	Asp	Arg	Phe	Thr	Gly	Ser	Gly	Ser	Gly	Thr	Asp	Phe	Thr
				85					90					95	

Leu	Lys	Ile	Ser	Arg	Val	Glu	Ala	Glu	Asp	Leu	Gly	Val	Tyr	Phe	Cys
			100					105					110		

Cys	Gln	Gly	Thr	His	Phe	Pro	Trp	Thr	Phe	Gly	Gly	Gly	Thr	Met	Leu
		115					120					125			

Glu	Ile	Lys
		130

<210> 1070  
 <211> 69  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 28F9 light chain variable framework 1 (FW1)  
 sequence

<400> 1070

gatgttgtga tgacccagac tccactcact ttgtcggta ccattggaca accagcctcc 60

atctcttgc 69

<210> 1071

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 28F9 light chain variable framework 1 (FW1) sequence

<400> 1071

Asp Val Val Met Thr Gln Thr Pro Leu Thr Leu Ser Val Thr Ile Gly  
1 5 10 15

Gln Pro Ala Ser Ile Ser Cys  
20

<210> 1072

<211> 48

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 28F9 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1072

aagtcaagtc agagcctctt acatagtgat ggaaagacat atttgatt 48

<210> 1073

<211> 16

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 28F9 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1073

Lys Ser Ser Gln Ser Leu Leu His Ser Asp Gly Lys Thr Tyr Leu Ile  
1 5 10 15

<210> 1074

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 28F9 light chain variable framework 2 (FW2) sequence

<400> 1074

tggttggttac agaggccagg ccagtcctca aagcgcctaa tctat 45

<210> 1075

<211> 15

<212> PRT

<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 light chain variable framework 2 (FW2)  
sequence

<400> 1075

Trp Leu Leu Gln Arg Pro Gly Gln Ser Pro Lys Arg Leu Ile Tyr  
1 5 10 15

<210> 1076  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1076  
ctggtgtcta aactggactc t 21

<210> 1077  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1077

Leu Val Ser Lys Leu Asp Ser  
1 5

<210> 1078  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 light chain variable framework 3 (FW3)  
sequence

<400> 1078  
ggagtccctg acaggttcac cggcagtgga tcaggacag atttcacact gaaaatcagc 60

agagtggagg ctgaggattt gggagtttat ttttgc 96

<210> 1079  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 light chain variable framework 3 (FW3)  
sequence

<400> 1079

Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys  
20 25 30

<210> 1080  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1080  
tgtcaaggta cacattttcc gtggacg 27

<210> 1081  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1081

Cys Gln Gly Thr His Phe Pro Trp Thr  
1 5

<210> 1082  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 light chain variable framework 4 (FW4)  
sequence

<400> 1082  
ttcgggtggag gcaccatgct ggaaatcaaa 30

<210> 1083  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 28F9 light chain variable framework 4 (FW4)  
sequence

<400> 1083

Phe Gly Gly Gly Thr Met Leu Glu Ile Lys  
1 5 10

<210> 1084  
<211> 411  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 Heavy chain: DNA sequence (411 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1084  
atgtacttgg gactgaacta tgtattcata gtttttctct taaatggtgt ccagagtga 60  
gtgaaacttg aggagtctgg aggaggcttg gtgcaacctg ggggatccat gaaactctct 120  
tgtgtgcct ctggattcac ttttaatgac gcctggatgg actgggtccg ccagtctcca 180  
gagaaggggc ttgagtgggt tgctgaaatt agaagcacag ctaatattca tacaacatac 240  
tatgctgagt ctgtccaagg gaggttcacc atctcaagag atgattcaa aagtagtgc 300  
tacctgcaa tgaacagctt gagagctgaa gacactggca tttattattg taccaccatta 360  
ctctacggat ttgcttactg gggccaaggg actctggtca ctgtctctgc a 411

<210> 1085

<211> 137

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 Heavy chain: Amino acid sequence (137 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1085

Met Tyr Leu Gly Leu Asn Tyr Val Phe Ile Val Phe Leu Leu Asn Gly  
1 5 10 15

Val Gln Ser Glu Val Lys Leu Glu Glu Ser Gly Gly Gly Leu Val Gln  
20 25 30

Pro Gly Gly Ser Met Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe  
35 40 45

Asn Asp Ala Trp Met Asp Trp Val Arg Gln Ser Pro Glu Lys Gly Leu  
50 55 60

Glu Trp Val Ala Glu Ile Arg Ser Thr Ala Asn Ile His Thr Thr Tyr  
65 70 75 80

Tyr Ala Glu Ser Val Gln Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser  
85 90 95

Lys Ser Ser Val Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110

Gly Ile Tyr Tyr Cys Thr Pro Leu Leu Tyr Gly Phe Ala Tyr Trp Gly  
115 120 125

Gln Gly Thr Leu Val Thr Val Ser Ala  
130 135

<210> 1086

<211> 90

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 heavy chain variable framework 1 (FW1)  
sequence

<400> 1086  
gaagtgaac ttgaggagtc tggaggaggc ttggtgcaac ctgggggatc catgaaactc 60  
tcttgctgctg cctctggatt cacttttaat 90

<210> 1087  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 heavy chain variable framework 1 (FW1)  
sequence

<400> 1087  
Glu Val Lys Leu Glu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly  
1 5 10 15

Ser Met Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn  
20 25 30

<210> 1088  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1088  
gacgcctgga tggac 15

<210> 1089  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1089  
Asp Ala Trp Met Asp  
1 5

<210> 1090  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 heavy chain variable framework 2 (FW2)  
sequence

<400> 1090  
tgggtccgcc agtctccaga gaaggggctt gagtgggttg ct 42

<210> 1091  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 heavy chain variable framework 2 (FW2)  
sequence

<400> 1091

Trp Val Arg Gln Ser Pro Glu Lys Gly Leu Glu Trp Val Ala  
1 5 10

<210> 1092  
<211> 57  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1092  
gaaattagaa gcacagctaa tattcatata acatactatg ctgagtctgt ccaaggg 57

<210> 1093  
<211> 19  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1093

Glu Ile Arg Ser Thr Ala Asn Ile His Thr Thr Tyr Tyr Ala Glu Ser  
1 5 10 15

Val Gln Gly

<210> 1094  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 heavy chain variable framework 3 (FW3)  
sequence

<400> 1094  
agggtcacca tctcaagaga tgattccaaa agtagtgtct acctgcaaat gaacagcttg 60

agagctgaag acaactggcat ttattattgt acccca 96

<210> 1095  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 heavy chain variable framework 3 (FW3)



sequence

<400> 1095

Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Ser Ser Val Tyr Leu Gln  
1 5 10 15

Met Asn Ser Leu Arg Ala Glu Asp Thr Gly Ile Tyr Tyr Cys Thr Pro  
20 25 30

<210> 1096

<211> 21

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1096

ttactctacg gatttgctta c

21

<210> 1097

<211> 7

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1097

Leu Leu Tyr Gly Phe Ala Tyr  
1 5

<210> 1098

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 heavy chain variable framework 4 (FW4)  
sequence

<400> 1098

tggggccaag ggactctggt cactgtctct gca

33

<210> 1099

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 heavy chain variable framework 4 (FW4)  
sequence

<400> 1099

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
1 5 10

<210> 1100

<211> 393  
<212> DNA  
<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 Light chain: DNA sequence (393 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1100

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atgaagttgc ctgttaggct gttggtgctg atgttctgga ttcctgcttc cagcagtgat      60
gttgtgatga cccaaagtcc actctccctg cctgtcagtc ttggagatca agcctccatc      120
tcttgcagaa ctagtcagag ccttgtacac agtaatggaa acacctatTT acattggcac      180
ctgcagaagc caggccagtc tccaaaggtc ctgatctaca aagtttccag ccgattttct      240
gggggtcccag acaggttcag tggcagtgga tcggggacag atttcacact caagatcagc      300
agagtggagg ctgaggatct gggagtttat ttctgctctc aaaatacaca tgttccgtac      360
acgttcggag gggggaccaa gctggaaata aaa                                393
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<210> 1101

<211> 131

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 Light chain: Amino acid sequence (131 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1101

```
Met Lys Leu Pro Val Arg Leu Leu Val Leu Met Phe Trp Ile Pro Ala
1              5              10             15
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```
Ser Ser Ser Asp Val Val Met Thr Gln Ser Pro Leu Ser Leu Pro Val
          20             25             30
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```
Ser Leu Gly Asp Gln Ala Ser Ile Ser Cys Arg Thr Ser Gln Ser Leu
          35             40             45
```

```
Val His Ser Asn Gly Asn Thr Tyr Leu His Trp His Leu Gln Lys Pro
          50             55             60
```

```
Gly Gln Ser Pro Lys Val Leu Ile Tyr Lys Val Ser Ser Arg Phe Ser
65             70             75             80
```

```
Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr
          85             90             95
```

```
Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys
          100            105            110
```

```
Ser Gln Asn Thr His Val Pro Tyr Thr Phe Gly Gly Gly Thr Lys Leu
          115            120            125
```

```
Glu Ile Lys
          130
```

<210> 1102  
<211> 69  
<212> DNA  
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<220>  
<223> Mouse antibody 18B4 light chain variable framework 1 (FW1) sequence

<400> 1102  
gatgttgta tgacccaaag tccactctcc ctgcctgtca gtcttggaga tcaagcctcc 60  
atctcttgc 69

<210> 1103  
<211> 23  
<212> PRT  
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<220>  
<223> Mouse antibody 18B4 light chain variable framework 1 (FW1) sequence

<400> 1103

Asp	Val	Val	Met	Thr	Gln	Ser	Pro	Leu	Ser	Leu	Pro	Val	Ser	Leu	Gly
1				5				10					15		

Asp Gln Ala Ser Ile Ser Cys  
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<210> 1104  
<211> 48  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1104  
agaactagtc agagccttgt acacagtaat ggaaacacct atttacat 48

<210> 1105  
<211> 16  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 18B4 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1105

Arg	Thr	Ser	Gln	Ser	Leu	Val	His	Ser	Asn	Gly	Asn	Thr	Tyr	Leu	His
1			5					10					15		

<210> 1106  
<211> 45  
<212> DNA  
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<220>  
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 <400> 1106  
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 <210> 1107  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 18B4 light chain variable framework 2 (FW2) sequence  
  
 <400> 1107  
  
 Trp His Leu Gln Lys Pro Gly Gln Ser Pro Lys Val Leu Ile Tyr  
 1 5 10 15  
  
 <210> 1108  
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 <212> DNA  
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 <223> Mouse antibody 18B4 light chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 1108  
 aaagtttcca gccgattttc t 21  
  
 <210> 1109  
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 <212> PRT  
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 <223> Mouse antibody 18B4 light chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 1109  
  
 Lys Val Ser Ser Arg Phe Ser  
 1 5  
  
 <210> 1110  
 <211> 96  
 <212> DNA  
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 <223> Mouse antibody 18B4 light chain variable framework 3 (FW3) sequence  
  
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 agagtggagg ctgaggatct gggagtttat ttctgc 96  
  
 <210> 1111  
 <211> 32  
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<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 light chain variable framework 3 (FW3) sequence

<400> 1111

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys  
20 25 30

<210> 1112

<211> 27

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1112

tctcaaaata cacatgttcc gtacacg

27

<210> 1113

<211> 9

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1113

Ser Gln Asn Thr His Val Pro Tyr Thr  
1 5

<210> 1114

<211> 30

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 18B4 light chain variable framework 4 (FW4) sequence

<400> 1114

ttcggagggg ggaccaagct ggaaataaaa

30

<210> 1115

<211> 10

<212> PRT

<213> Artificial Sequence

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<223> Mouse antibody 18B4 light chain variable framework 4 (FW4) sequence

<400> 1115

Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys

1 5 10

<210> 1116  
<211> 408  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse Antibody IE4 Heavy chain: DNA sequence (408 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1116  
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gttcagctgc agcagtctgg ggctgagctg gtgaggcctg ggtcctcagt gaagatttcc 120  
tgtaaggctt ctggctatgc attcagtacc tactggatga actgggtgaa gcagaggcct 180  
ggacagggtc ttgagtggat tggacagatt tatcctggag atagtgatac taactacaat 240  
ggaaaattca agggtaaagc cacactgact gcagacaagt cctccaacac agcctacatg 300  
cagctcagca gcctaacatc tgaggactct gcggtctttt tctgtgcaag aggtaaccac 360  
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<210> 1117  
<211> 136  
<212> PRT  
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<220>  
<223> Mouse Antibody IE4 Heavy chain: Amino acid sequence (136 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1117

Met Glu Trp Pro Cys Ile Phe Leu Phe Leu Leu Ser Val Thr Glu Gly  
1 5 10 15

Val His Ser Gln Val Gln Leu Gln Gln Ser Gly Ala Glu Leu Val Arg  
20 25 30

Pro Gly Ser Ser Val Lys Ile Ser Cys Lys Ala Ser Gly Tyr Ala Phe  
35 40 45

Ser Thr Tyr Trp Met Asn Trp Val Lys Gln Arg Pro Gly Gln Gly Leu  
50 55 60

Glu Trp Ile Gly Gln Ile Tyr Pro Gly Asp Ser Asp Thr Asn Tyr Asn  
65 70 75 80

Gly Lys Phe Lys Gly Lys Ala Thr Leu Thr Ala Asp Lys Ser Ser Asn  
85 90 95

Thr Ala Tyr Met Gln Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val  
100 105 110

Phe Phe Cys Ala Arg Gly Asn His Ala Ser Met Asp Tyr Trp Gly Gln  
115 120 125

Gly Thr Ser Val Thr Val Ser Ser  
130 135

<210> 1118  
<211> 90  
<212> DNA  
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<220>  
<223> Mouse Antibody IE4 heavy chain variable framework 1 (FW1)  
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<400> 1118  
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tcctgtaagg cttctggcta tgcattcagt 90

<210> 1119  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse Antibody IE4 heavy chain variable framework 1 (FW1)  
sequence

<400> 1119

Gln Val Gln Leu Gln Gln Ser Gly Ala Glu Leu Val Arg Pro Gly Ser  
1 5 10 15

Ser Val Lys Ile Ser Cys Lys Ala Ser Gly Tyr Ala Phe Ser  
20 25 30

<210> 1120

<400> 1120  
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<210> 1200

<211> 15

<212> DNA

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<220>

<223> Mouse antibody IE4 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1200

acctactgga tgaac

15

<210> 1201

<211> 5

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody IE4 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1201

Thr Tyr Trp Met Asn

1 5

<210> 1202

<211> 42

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody IE4 heavy chain variable framework 2 (FW2)  
sequence

<400> 1202

tgggtgaagc agaggcctgg acagggtctt gagtggattg ga

42

<210> 1203

<211> 14

<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 heavy chain variable framework 2 (FW2)  
sequence

<400> 1203

Trp Val Lys Gln Arg Pro Gly Gln Gly Leu Glu Trp Ile Gly  
1 5 10

<210> 1204  
<211> 51  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1204  
cagatattatc ctggagatag tgatactaac tacaatggaa agttcaaggg t 51

<210> 1205  
<211> 17  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1205

Gln Ile Tyr Pro Gly Asp Ser Asp Thr Asn Tyr Asn Gly Lys Phe Lys  
1 5 10 15

Gly

<210> 1206  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 heavy chain variable framework 3 (FW3)  
sequence

<400> 1206  
aaagccacac tgactgcaga caagtcctcc aacacagcct acatgcagct cagcagccta 60

acatctgagg actctgcggt ctttttctgt gcaaga 96

<210> 1207  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 heavy chain variable framework 3 (FW3)  
sequence

<400> 1207

Lys Ala Thr Leu Thr Ala Asp Lys Ser Ser Asn Thr Ala Tyr Met Gln  
1 5 10 15

Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val Phe Phe Cys Ala Arg  
20 25 30

<210> 1208

<211> 24

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody IE4 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1208

ggtaaccacg cctctatgga ctac

24

<210> 1209

<211> 8

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody IE4 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1209

Gly Asn His Ala Ser Met Asp Tyr  
1 5

<210> 1210

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody IE4 heavy chain variable framework 4 (FW4)  
sequence

<400> 1210

tgggggtcaag gaacctcagt caccgtctcc tca

33

<210> 1211

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody IE4 heavy chain variable framework 4 (FW4)  
sequence

<400> 1211

Trp Gly Gln Gly Thr Ser Val Thr Val Ser Ser  
1 5 10

<210> 1212

<211> 393

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse Antibody IE4 Light chain: DNA sequence (393 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1212

```
atgaagttgc ctgttaggct gttggtgctg atgttctgga ttcctgcttc cagcagtgat      60
gttgtgatga cccaaactcc actctccctg cctgtcagtc ttggagatca agcctccatc      120
tcttgcagat ctagtcagag ccttgtagac agtaatggaa acacctatct acattggtac      180
ctgcagaagc caggccagtc tccaaagctc ctgatctaca aagtttccaa ccgattttct      240
ggggtcccag acaggttcag tggcagtggg tcagggacag atttcacact caagatcagc      300
agagtggagg ctgaggatct gggagtttat ttctgctctc aaaaaacaca tgttccgtgg      360
acgttcggtg gaggcaccaa gctggaaatc aaa                                  393
```

<210> 1213

<211> 131

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse Antibody IE4 Light chain: Amino acid sequence (131 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1213

```
Met Lys Leu Pro Val Arg Leu Leu Val Leu Met Phe Trp Ile Pro Ala
1              5              10             15
```

```
Ser Ser Ser Asp Val Val Met Thr Gln Thr Pro Leu Ser Leu Pro Val
          20             25             30
```

```
Ser Leu Gly Asp Gln Ala Ser Ile Ser Cys Arg Ser Ser Gln Ser Leu
          35             40             45
```

```
Val His Ser Asn Gly Asn Thr Tyr Leu His Trp Tyr Leu Gln Lys Pro
          50             55             60
```

```
Gly Gln Ser Pro Lys Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser
          65             70             75             80
```

```
Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr
          85             90             95
```

```
Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys
          100            105            110
```

```
Ser Gln Lys Thr His Val Pro Trp Thr Phe Gly Gly Gly Thr Lys Leu
          115            120            125
```

```
Glu Ile Lys
          130
```



<210> 1214  
 <211> 69  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody IE4 light chain variable framework 1 (FW1) sequence  
  
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 gatgttgta tgacccaaac tccactctcc ctgcctgtca gtcttggaga tcaagcctcc 60  
 atctcttgc 69

<210> 1215  
 <211> 23  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody IE4 light chain variable framework 1 (FW1) sequence  
  
 <400> 1215  
  
 Asp Val Val Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
 1 5 10 15  
  
 Asp Gln Ala Ser Ile Ser Cys  
 20

<210> 1216  
 <211> 48  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody IE4 light chain variable complementarity determining regions 1 (CDR1) sequence  
  
 <400> 1216  
 agatctagtc agagccttgt acacagtaat ggaaacacct atttacat 48

<210> 1217  
 <211> 16  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody IE4 light chain variable complementarity determining regions 1 (CDR1) sequence  
  
 <400> 1217  
  
 Arg Ser Ser Gln Ser Leu Val His Ser Asn Gly Asn Thr Tyr Leu His  
 1 5 10 15

<210> 1218  
 <211> 45  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody IE4 light chain variable framework 2 (FW2)

# sequence

<400> 1218  
 tggtagcctgc agaagccagg ccagtctcca aagctcctga tctac 45

<210> 1219  
 <211> 15  
 <212> PRT  
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<220>  
 <223> Mouse antibody IE4 light chain variable framework 2 (FW2)  
 sequence

<400> 1219  
 Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
 1 5 10 15

<210> 1220  
 <211> 21  
 <212> DNA  
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<220>  
 <223> Mouse antibody IE4 light chain variable complementarity  
 determining regions 2 (CDR2) sequence

<400> 1220  
 aaagtttcca accgattttc t 21

<210> 1221  
 <211> 7  
 <212> PRT  
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<220>  
 <223> Mouse antibody IE4 light chain variable complementarity  
 determining regions 2 (CDR2) sequence

<400> 1221  
 Lys Val Ser Asn Arg Phe Ser  
 1 5

<210> 1222  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody IE4 light chain variable framework 3 (FW3)  
 sequence

<400> 1222  
 ggggtccag acaggttcag tggcagtgga tcagggacag atttcacact caagatcagc 60

agagtggagg ctgaggatct gggagtttat ttctgc 96

<210> 1223  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 light chain variable framework 3 (FW3)  
sequence

<400> 1223

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys  
20 25 30

<210> 1224  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1224

tctcaaaaaa cacatgttcc gtggacg

27

<210> 1225  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1225

Ser Gln Lys Thr His Val Pro Trp Thr  
1 5

<210> 1226  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 light chain variable framework 4 (FW4)  
sequence

<400> 1226

ttcgggtggag gcaccaagct ggaaatcaaa

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<210> 1227  
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<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody IE4 light chain variable framework 4 (FW4)  
sequence

<400> 1227

Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
1 5 10

<210> 1228  
 <211> 411  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 29H1 Heavy chain: DNA sequence (411 bp) - Signal  
 sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1228  
 atgtacttgg gactgaacta tgtattcata gtttttctct taaatggtgt ccagagtgaa 60  
 gtgaagcttg aggagtctgg aggaggcttg gtacaacctg gaggatccat gaaactctct 120  
 tgtgctgcct ctggattcac ttttagtgac gcctggatgg actgggtccg ccagtctcca 180  
 gagaaggggc ttgaatgggt tgctgaaatt agaagcaaag ctactaatca tgcaacatac 240  
 tatgctgagt ctgtgaaagg gaggttcacc atctcaagag atgattccaa aagtagtgtc 300  
 tacctgcaaa tgaacagctt aagagctgaa gacactggca tttattactg taccctcccta 360  
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<210> 1229  
 <211> 137  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 29H1 Heavy chain: Amino acid sequence (137 aa) -  
 Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1229  
 Met Tyr Leu Gly Leu Asn Tyr Val Phe Ile Val Phe Leu Leu Asn Gly  
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 Val Gln Ser Glu Val Lys Leu Glu Glu Ser Gly Gly Gly Leu Val Gln  
 20 25 30  
 Pro Gly Gly Ser Met Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe  
 35 40 45  
 Ser Asp Ala Trp Met Asp Trp Val Arg Gln Ser Pro Glu Lys Gly Leu  
 50 55 60  
 Glu Trp Val Ala Glu Ile Arg Ser Lys Ala Thr Asn His Ala Thr Tyr  
 65 70 75 80  
 Tyr Ala Glu Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser  
 85 90 95  
 Lys Ser Ser Val Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
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 Gly Ile Tyr Tyr Cys Thr Pro Leu Leu Tyr Gly Phe Ala Tyr Trp Gly  
 115 120 125

Gln Gly Thr Leu Val Thr Val Ser Ala  
130 135

<210> 1230  
<211> 90  
<212> DNA  
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<220>  
<223> Mouse antibody 29H1 heavy chain variable framework 1 (FW1)  
sequence

<400> 1230  
gaagtgaagc ttgaggagtc tggaggaggc ttggtacaac ctggaggatc catgaaactc 60  
tcttgtgctg cctctggatt cacttttagt 90

<210> 1231  
<211> 30  
<212> PRT  
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<220>  
<223> Mouse antibody 29H1 heavy chain variable framework 1 (FW1)  
sequence

<400> 1231

Glu Val Lys Leu Glu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly  
1 5 10 15

Ser Met Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 1232  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1232  
gacgcctgga tggac 15

<210> 1233  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1233

Asp Ala Trp Met Asp  
1 5

<210> 1234  
<211> 42  
<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable framework 2 (FW2) sequence

<400> 1234

tgggtccgcc agtctccaga gaaggggctt gaatgggttg ct

42

<210> 1235

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable framework 2 (FW2) sequence

<400> 1235

Trp Val Arg Gln Ser Pro Glu Lys Gly Leu Glu Trp Val Ala

1 5 10

<210> 1236

<211> 57

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1236

gaaattagaa gcaaagctac taatcatgca acatactatg ctgagtctgt gaaaggg

57

<210> 1237

<211> 19

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1237

Glu Ile Arg Ser Lys Ala Thr Asn His Ala Thr Tyr Tyr Ala Glu Ser

1 5 10 15

Val Lys Gly

<210> 1238

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable framework 3 (FW3) sequence

<400> 1238

aggttcacca tctcaagaga tgattccaaa agtagtgtct acctgcaaat gaacagctta

60

agagctgaag acactggcat ttattactgt accccc

96

<210> 1239

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable framework 3 (FW3) sequence

<400> 1239

Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Ser Ser Val Tyr Leu Gln  
1 5 10 15

Met Asn Ser Leu Arg Ala Glu Asp Thr Gly Ile Tyr Tyr Cys Thr Pro  
20 25 30

<210> 1240

<211> 21

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1240

ctactttacg ggtttgctta c

21

<210> 1241

<211> 7

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1241

Leu Leu Tyr Gly Phe Ala Tyr  
1 5

<210> 1242

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable framework 4 (FW4) sequence

<400> 1242

tggggccaag ggactctggc cactgtctct gca

33

<210> 1243

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 heavy chain variable framework 4 (FW4)  
sequence

<400> 1243

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
1 5 10

<210> 1244

<211> 393

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 Light chain: DNA sequence (393 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1244

atgaagtgc ctgttaggct gttggtgctg atgttctgga ttcctgcttc cagcagtgat 60  
gttgtgatga cccaaactcc actctccctg cctgtcagtc ttggagatca agcctccatc 120  
tcttcagat ctggtcagag ccttgtagac agtaatggac acacctatct acattggtac 180  
ctgcagaagc caggccagtc tccaaggctc ctgatctaca aagtttccaa ccgattttct 240  
gggggtcccag acaggttcag tggcagtgga tcaagggcag atttcacact caagatcagc 300  
agagtggagg ctgaggatct gggagtttat ttctgctctc aaactacaca tgttccgtgg 360  
acgttcggtg gaggcaccaa gctggaaatc aaa 393

<210> 1245

<211> 131

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 29H1 Light chain: Amino acid sequence (131 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1245

Met Lys Leu Pro Val Arg Leu Leu Val Leu Met Phe Trp Ile Pro Ala  
1 5 10 15

Ser Ser Ser Asp Val Val Met Thr Gln Thr Pro Leu Ser Leu Pro Val  
20 25 30

Ser Leu Gly Asp Gln Ala Ser Ile Ser Cys Arg Ser Gly Gln Ser Leu  
35 40 45

Val His Ser Asn Gly His Thr Tyr Leu His Trp Tyr Leu Gln Lys Pro  
50 55 60

Gly Gln Ser Pro Arg Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser  
65 70 75 80

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Arg Ala Asp Phe Thr  
85 90 95



Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys  
100 105 110

Ser Gln Thr Thr His Val Pro Trp Thr Phe Gly Gly Gly Thr Lys Leu  
115 120 125

Glu Ile Lys  
130

<210> 1246  
<211> 69  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable framework 1 (FW1)  
sequence

<400> 1246  
gatgttgtaga tgacccaaac tccactctcc ctgcctgtca gtcttgaga tcaagcctcc 60  
atctcttgc 69

<210> 1247  
<211> 23  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable framework 1 (FW1)  
sequence

<400> 1247

Asp Val Val Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
1 5 10 15

Asp Gln Ala Ser Ile Ser Cys  
20

<210> 1248  
<211> 48  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1248  
agatctgggtc agagccttgt acacagtaat ggacacacct atttacat 48

<210> 1249  
<211> 16  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1249

Arg Ser Gly Gln Ser Leu Val His Ser Asn Gly His Thr Tyr Leu His  
1 5 10 15

<210> 1250  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable framework 2 (FW2)  
sequence

<400> 1250  
tggtagctgc agaagccagg ccagtctcca aggtctctga tctac 45

<210> 1251  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable framework 2 (FW2)  
sequence

<400> 1251

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr  
1 5 10 15

<210> 1252  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1252  
aaagtttcca accgattttc t 21

<210> 1253  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1253

Lys Val Ser Asn Arg Phe Ser  
1 5

<210> 1254  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable framework 3 (FW3)  
sequence

<400> 1254  
ggggtcccag acaggttcag tggcagtgga tcaagggcag atttcacact caagatcagc 60  
agagtggagg ctgaggatct gggagtttat ttctgc 96

<210> 1255  
<211> 32  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 29H1 light chain variable framework 3 (FW3)  
sequence

<400> 1255  
Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Arg Ala Asp Phe Thr  
1 5 10 15  
Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys  
20 25 30

<210> 1256  
<211> 27  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 29H1 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1256  
tctcaaaact cacatgttcc gtggacg 27

<210> 1257  
<211> 9  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 29H1 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1257  
Ser Gln Thr Thr His Val Pro Trp Thr  
1 5

<210> 1258  
<211> 30  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 29H1 light chain variable framework 4 (FW4)  
sequence

<400> 1258  
ttcggtaggag gcaccaagct ggaaatcaaa 30

<210> 1259  
<211> 10

<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 29H1 light chain variable framework 4 (FW4) sequence

<400> 1259

Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
1 5 10

<210> 1260  
<211> 399  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 Heavy chain: DNA sequence (399 bp) - Signal sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1260  
atggaaggc actggatctt tctcttcctg ttttcagtaa ctgcagggtg ccactcccag 60  
gtccagcttc agcagtctgg ggctgaactg gcaaaacctg gggcctcagt gaagatgtcc 120  
tgcaaggctt ctggctacac ctttactagc tactggatgc actgggtaaa acagaggcct 180  
ggacagggtc tggaatggat tggatacatt aatcctagca ctggttatac tgagtacaat 240  
cagaagttca aggacaaggc cacattgact gcagacaaat cctccagcac agcctacatg 300  
caactgagca gcctgacatc tgaggactct gcagtctatt actgtgcaag agcctacatt 360  
gactactggg gccaaggcac cactctcaca gtctcctca 399

<210> 1261  
<211> 133  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 Heavy chain: Amino acid sequence (133 aa) - Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1261

Met Glu Arg His Trp Ile Phe Leu Phe Leu Phe Ser Val Thr Ala Gly  
1 5 10 15

Val His Ser Gln Val Gln Leu Gln Gln Ser Gly Ala Glu Leu Ala Lys  
20 25 30

Pro Gly Ala Ser Val Lys Met Ser Cys Lys Ala Ser Gly Tyr Thr Phe  
35 40 45

Thr Ser Tyr Trp Met His Trp Val Lys Gln Arg Pro Gly Gln Gly Leu  
50 55 60

Glu Trp Ile Gly Tyr Ile Asn Pro Ser Thr Gly Tyr Thr Glu Tyr Asn  
65 70 75 80

Gln Lys Phe Lys Asp Lys Ala Thr Leu Thr Ala Asp Lys Ser Ser Ser  
85 90 95

Thr Ala Tyr Met Gln Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val  
100 105 110

Tyr Tyr Cys Ala Arg Ala Tyr Ile Asp Tyr Trp Gly Gln Gly Thr Thr  
115 120 125

Leu Thr Val Ser Ser  
130

<210> 1262  
<211> 90  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 heavy chain variable framework 1 (FW1) sequence

<400> 1262  
cagggtccagc ttcagcagtc tggggctgaa ctggcaaaac ctggggcctc agtgaagatg 60  
tcctgcaagg cttctggcta cacctttact 90

<210> 1263  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 heavy chain variable framework 1 (FW1) sequence

<400> 1263

Gln Val Gln Leu Gln Gln Ser Gly Ala Glu Leu Ala Lys Pro Gly Ala  
1 5 10 15

Ser Val Lys Met Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr  
20 25 30

<210> 1264  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1264  
agctactgga tgcac 15

<210> 1265  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>

<223> Mouse antibody 31A1 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1265

Ser Tyr Trp Met His  
1 5

<210> 1266

<211> 42

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 31A1 heavy chain variable framework 2 (FW2)  
sequence

<400> 1266

tgggtaaac agaggcctgg acagggtctg gaatggattg ga

42

<210> 1267

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 31A1 heavy chain variable framework 2 (FW2)  
sequence

<400> 1267

Trp Val Lys Gln Arg Pro Gly Gln Gly Leu Glu Trp Ile Gly  
1 5 10

<210> 1268

<211> 51

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 31A1 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1268

tacattaatc ctagcactgg ttatactgag tacaatcaga agttcaagga c

51

<210> 1269

<211> 17

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 31A1 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1269

Tyr Ile Asn Pro Ser Thr Gly Tyr Thr Glu Tyr Asn Gln Lys Phe Lys  
1 5 10 15

Asp

<210> 1270  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 31A1 heavy chain variable framework 3 (FW3) sequence  
  
 <400> 1270  
 aaggccacat tgactgcaga caaatcctcc agcacagcct acatgcaact gagcagcctg 60  
 acatctgagg actctgcagt ctattactgt gcaaga 96

<210> 1271  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 31A1 heavy chain variable framework 3 (FW3) sequence  
  
 <400> 1271  
  
 Lys Ala Thr Leu Thr Ala Asp Lys Ser Ser Ser Thr Ala Tyr Met Gln  
 1 5 10 15

Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val Tyr Tyr Cys Ala Arg  
 20 25 30

<210> 1272  
 <211> 15  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 31A1 heavy chain variable complementarity determining regions 3 (CDR3) sequence  
  
 <400> 1272  
 gcctacattg actac 15

<210> 1273  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 31A1 heavy chain variable complementarity determining regions 3 (CDR3) sequence  
  
 <400> 1273  
  
 Ala Tyr Ile Asp Tyr  
 1 5

<210> 1274  
 <211> 33  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 31A1 heavy chain variable framework 4 (FW4)

# sequence

<400> 1274  
tggggccaag gcaccactct cacagtctcc tca 33

<210> 1275  
<211> 11  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 heavy chain variable framework 4 (FW4)  
sequence

<400> 1275  
Trp Gly Gln Gly Thr Thr Leu Thr Val Ser Ser  
1 5 10

<210> 1276  
<211> 393  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 Light chain: DNA sequence (393 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1276  
atgaagttgc ctgttaggct gttggtgctg atgttctgga ttcctgcttc cagcagtgat 60  
gttttgatga cccaaactcc actctccctg cctgtcagtc ttggagatca agcctccttc 120  
tcttgcagat ctagtcagag cattgtacat agtaatggaa acacctatctt agaatggtac 180  
ctgcagaaac caggccagtc tccaaagctc ctgatctaca aagtttccaa ccgattttct 240  
ggggtcccag acaggtttcag tggcagtgga tcaggacag atttcacact caagatcaac 300  
agagtggagg ctgaggatct gggagtttat tactgctttc aagtttcaca ttttccgtgg 360  
acgttcggtg gaggcaccaa gctggaaatc aaa 393

<210> 1277  
<211> 131  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 Light chain: Amino acid sequence (131 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1277  
Met Lys Leu Pro Val Arg Leu Leu Val Leu Met Phe Trp Ile Pro Ala  
1 5 10 15

Ser Ser Ser Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val  
20 25 30

Ser Leu Gly Asp Gln Ala Ser Phe Ser Cys Arg Ser Ser Gln Ser Ile  
35 40 45



Val His Ser Asn Gly Asn Thr Tyr Leu Glu Trp Tyr Leu Gln Lys Pro  
50 55 60

Gly Gln Ser Pro Lys Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser  
65 70 75 80

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
85 90 95

Leu Lys Ile Asn Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
100 105 110

Phe Gln Val Ser His Phe Pro Trp Thr Phe Gly Gly Gly Thr Lys Leu  
115 120 125

Glu Ile Lys  
130

<210> 1278  
<211> 69  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable framework 1 (FW1)  
sequence

<400> 1278  
gatgttttga tgacccaaac tccactctcc ctgcctgtca gtcttggaga tcaagcctcc 60  
ttctcttgc 69

<210> 1279  
<211> 23  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable framework 1 (FW1)  
sequence

<400> 1279

Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
1 5 10 15

Asp Gln Ala Ser Phe Ser Cys  
20

<210> 1280  
<211> 48  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1280  
agatctagtc agagcattgt acatagaaat ggaaacacct atttagaa 48

<210> 1281  
<211> 16  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1281

Arg Ser Ser Gln Ser Ile Val His Arg Asn Gly Asn Thr Tyr Leu Glu  
1 5 10 15

<210> 1282  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable framework 2 (FW2)  
sequence

<400> 1282  
tggtagctgc agaaaccagg ccagtctcca aagctcctga tctac 45

<210> 1283  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable framework 2 (FW2)  
sequence

<400> 1283

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 1284  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1284  
aaagtttcca accgattttc t 21

<210> 1285  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1285

Lys Val Ser Asn Arg Phe Ser  
1 5

<210> 1286  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable framework 3 (FW3)  
sequence

<400> 1286  
ggggtcccag acaggttcag tggcagtgga tcagggacag atttcacact caagatcaac 60  
agagtggagg ctgaggatct gggagtttat tactgc 96

<210> 1287  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable framework 3 (FW3)  
sequence

<400> 1287

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Lys Ile Asn Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
20 25 30

<210> 1288  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1288  
tttcaagttt cacattttcc gtggacg 27

<210> 1289  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 31A1 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1289

Phe Gln Val Ser His Phe Pro Trp Thr  
1 5

<210> 1290  
<211> 30  
<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 31A1 light chain variable framework 4 (FW4) sequence

<400> 1290

ttcgggtggag gcaccaagct ggaaatcaaa

30

<210> 1291

<211> 10

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 31A1 light chain variable framework 4 (FW4) sequence

<400> 1291

Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys

1 5 10

<210> 1292

<211> 411

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 32C1 Heavy chain: DNA sequence (411 bp) - Signal sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1292

atgtacttgg gactgaactg tgtattcata gtttttctct taaaaggtgt ccagagtgaa 60

gtgaagcttg aggagtcttg aggaggcttg gtgcaatctg gaggatccat gaaactctcc 120

tgtgttgctt ctggattcac tttcagtaat tactggatga actgggtccg ccagtctcca 180

gagaaggggc ttgagtgggt tgctgaaatt agattgaaat ctaataatta tgcaatacat 240

tatgcgagtg ctgtgaaggg gaggttcacc atctcaagag atgattcaa aagtagtgtc 300

tacctgcaaa tgaacaactt aagagctgaa gacactggca tttattactg taccagggtc 360

ccgggactgg atgcttactg gggccaaggg actctggtca ctgtctctgc a 411

<210> 1293

<211> 137

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 32C1 Heavy chain: Amino acid sequence (137 aa) - Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1293

Met Tyr Leu Gly Leu Asn Cys Val Phe Ile Val Phe Leu Leu Lys Gly

1 5 10 15

Val Gln Ser Glu Val Lys Leu Glu Glu Ser Gly Gly Gly Leu Val Gln

20 25 30

Ser Gly Gly Ser Met Lys Leu Ser Cys Val Ala Ser Gly Phe Thr Phe  
35 40 45

Ser Asn Tyr Trp Met Asn Trp Val Arg Gln Ser Pro Glu Lys Gly Leu  
50 55 60

Glu Trp Val Ala Glu Ile Arg Leu Lys Ser Asn Asn Tyr Ala Ile His  
65 70 75 80

Tyr Ala Glu Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser  
85 90 95

Lys Ser Ser Val Tyr Leu Gln Met Asn Asn Leu Arg Ala Glu Asp Thr  
100 105 110

Gly Ile Tyr Tyr Cys Thr Arg Val Pro Gly Leu Asp Ala Tyr Trp Gly  
115 120 125

Gln Gly Thr Leu Val Thr Val Ser Ala  
130 135

<210> 1294  
<211> 90  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable framework 1 (FW1)  
sequence

<400> 1294  
gaagtgaagc ttgaggagtc tggaggaggc ttggtgcaat ctggaggatc catgaaactc 60  
tcctgtgttg cctctggatt cactttcagt 90

<210> 1295  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable framework 1 (FW1)  
sequence

<400> 1295

Glu Val Lys Leu Glu Glu Ser Gly Gly Gly Leu Val Gln Ser Gly Gly  
1 5 10 15

Ser Met Lys Leu Ser Cys Val Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 1296  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable complementarity

determining regions 1 (CDR1) sequence

<400> 1296  
aattactgga tgaac 15

<210> 1297  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1297

Asn Tyr Trp Met Asn  
1 5

<210> 1298  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable framework 2 (FW2)  
sequence

<400> 1298  
tgggtccgcc agtctccaga gaaggggctt gagggttg ct 42

<210> 1299  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable framework 2 (FW2)  
sequence

<400> 1299

Trp Val Arg Gln Ser Pro Glu Lys Gly Leu Glu Trp Val Ala  
1 5 10

<210> 1300  
<211> 57  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1300  
gaaattagat tgaatctaa taattatgca atacattatg cggagtctgt gaagggg 57

<210> 1301  
<211> 19  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable complementarity

determining regions 2 (CDR2) sequence

<400> 1301

Glu Ile Arg Leu Lys Ser Asn Asn Tyr Ala Ile His Tyr Ala Glu Ser  
1 5 10 15

Val Lys Gly

<210> 1302

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 32C1 heavy chain variable framework 3 (FW3)  
sequence

<400> 1302

agggtcacca tctcaagaga tgattccaaa agtagtgtct acctgcaaat gaacaactta 60

agagctgaag aactggcat ttattactgt accagg 96

<210> 1303

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 32C1 heavy chain variable framework 3 (FW3)  
sequence

<400> 1303

Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Ser Ser Val Tyr Leu Gln  
1 5 10 15

Met Asn Asn Leu Arg Ala Glu Asp Thr Gly Ile Tyr Tyr Cys Thr Arg  
20 25 30

<210> 1304

<211> 21

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 32C1 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1304

gtcccgggac tggatgctta c 21

<210> 1305

<211> 7

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 32C1 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1305

Val Pro Gly Leu Asp Ala Tyr  
1 5

<210> 1306  
<211> 33  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable framework 4 (FW4)  
sequence

<400> 1306  
tggggccaag ggactctggt cactgtctct gca 33

<210> 1307  
<211> 11  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 heavy chain variable framework 4 (FW4)  
sequence

<400> 1307

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
1 5 10

<210> 1308  
<211> 393  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 Light chain: DNA sequence (393 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1308  
atgaagttgc ctgttaggct gttggtgctg atgttctgga ttcctgcttc cagcagtgat 60  
gttgtgatga cccaaactcc actctccctg cctgtcagtc ttggagatca agcctccatc 120  
tcttgcagat ctagtcagag ccttgtacac agtaatggaa acacctatctt acattggtac 180  
ctgcagaagc caggccagtc tccaaagctc ctgatctaca aagtttccaa ccgattttct 240  
gggggtcccag acaggttcag tggcagtgga tcaggacag atttcacact caagatcagc 300  
agtgtggagg ctgaggatct gggagtttat ttctgtcttc aaattacaca tgttccgtac 360  
acgttcggag gggggaccaa tctggaaata aaa 393

<210> 1309  
<211> 131  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 Light chain: Amino acid sequence (131 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1309



Met Lys Leu Pro Val Arg Leu Leu Val Leu Met Phe Trp Ile Pro Ala  
1 5 10 15

Ser Ser Ser Asp Val Val Met Thr Gln Thr Pro Leu Ser Leu Pro Val  
20 25 30

Ser Leu Gly Asp Gln Ala Ser Ile Ser Cys Arg Ser Ser Gln Ser Leu  
35 40 45

Val His Ser Asn Gly Asn Thr Tyr Leu His Trp Tyr Leu Gln Lys Pro  
50 55 60

Gly Gln Ser Pro Lys Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser  
65 70 75 80

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
85 90 95

Leu Lys Ile Ser Ser Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys  
100 105 110

Ser Gln Ile Thr His Val Pro Tyr Thr Phe Gly Gly Gly Thr Asn Leu  
115 120 125

Glu Ile Lys  
130

<210> 1310  
<211> 69  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 light chain variable framework 1 (FW1)  
sequence

<400> 1310  
gatgttgatga tgacccaaac tccactctcc ctgcctgtca gtcttgaga tcaagcctcc 60  
atctcttgc 69

<210> 1311  
<211> 23  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 32C1 light chain variable framework 1 (FW1)  
sequence

<400> 1311

Asp Val Val Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
1 5 10 15

Asp Gln Ala Ser Ile Ser Cys  
20

<210> 1312  
 <211> 48  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 32C1 light chain variable complementarity  
 determining regions 1 (CDR1) sequence  
  
 <400> 1312  
 agatctagtc agagccttgt acacagtaat ggaaacacct atttacat 48

<210> 1313  
 <211> 16  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 32C1 light chain variable complementarity  
 determining regions 1 (CDR1) sequence  
  
 <400> 1313  
  
 Arg Ser Ser Gln Ser Leu Val His Ser Asn Gly Asn Thr Tyr Leu His  
 1 5 10 15

<210> 1314  
 <211> 45  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 32C1 light chain variable framework 2 (FW2)  
 sequence  
  
 <400> 1314  
 tggtagctgc agaagccagg ccagtcctcca aagctcctga tctac 45

<210> 1315  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 32C1 light chain variable framework 2 (FW2)  
 sequence  
  
 <400> 1315  
  
 Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
 1 5 10 15

<210> 1316  
 <211> 21  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 32C1 light chain variable complementarity  
 determining regions 2 (CDR2) sequence  
  
 <400> 1316  
 aaagtttcca accgattttc t 21

<210> 1317  
 <211> 7  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 32C1 light chain variable complementarity  
 determining regions 2 (CDR2) sequence

<400> 1317

Lys Val Ser Asn Arg Phe Ser  
 1 5

<210> 1318  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 32C1 light chain variable framework 3 (FW3)  
 sequence

<400> 1318  
 ggggtcccag acaggttcag tggcagtgga tcaggacag atttcacact caagatcagc 60  
 agtgtggagg ctgaggatct gggagtttat ttctgc 96

<210> 1319  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 32C1 light chain variable framework 3 (FW3)  
 sequence

<400> 1319

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 1 5 10 15

Leu Lys Ile Ser Ser Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys  
 20 25 30

<210> 1320  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 32C1 light chain variable complementarity  
 determining regions 3 (CDR3) sequence

<400> 1320  
 tctcaaatta cacatgttcc gtacacg 27

<210> 1321  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence

<220>

<223> Mouse antibody 32C1 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1321

Ser Gln Ile Thr His Val Pro Tyr Thr  
1 5

<210> 1322

<211> 30

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 32C1 light chain variable framework 4 (FW4)  
sequence

<400> 1322

ttcggagggg ggaccaatct ggaaataaaa

30

<210> 1323

<211> 10

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 32C1 light chain variable framework 4 (FW4)  
sequence

<400> 1323

Phe Gly Gly Gly Thr Asn Leu Glu Ile Lys  
1 5 10

<210> 1324

<211> 423

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 Heavy chain: DNA sequence (423 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1324

atgaaatgca gctgggttat cttcttcctg atggcagtggt ttacaggggt caattcagag

60

gttcagctgc agcagtcctgg ggcagacctt gtgaagccag gggcctcagt caagttgtcc

120

tgcacagctt ctggcttcaa cattaaagac acctttatgc actgggtgaa gcagaggcct

180

gaacagggcc tggagtggat tggaaggatt gatcctgcga atggtaatac taaatatgac

240

ccgaaattcc agggcaaggc cactataaca gcagacacat cctccaacac agcctacctg

300

cagctcagca gcctgacatc tgaggacact gccgtctatt actgtgctaa accgtatggt

360

aactacggct attactatgc ttgggactac tgggggtcaag gaacctcagt caccgtctcc

420

tca

423

<210> 1325

<211> 141

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 Heavy chain: Amino acid sequence (141 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1325

Met Lys Cys Ser Trp Val Ile Phe Phe Leu Met Ala Val Val Thr Gly  
1 5 10 15

Val Asn Ser Glu Val Gln Leu Gln Gln Ser Gly Ala Asp Leu Val Lys  
20 25 30

Pro Gly Ala Ser Val Lys Leu Ser Cys Thr Ala Ser Gly Phe Asn Ile  
35 40 45

Lys Asp Thr Phe Met His Trp Val Lys Gln Arg Pro Glu Gln Gly Leu  
50 55 60

Glu Trp Ile Gly Arg Ile Asp Pro Ala Asn Gly Asn Thr Lys Tyr Asp  
65 70 75 80

Pro Lys Phe Gln Gly Lys Ala Thr Ile Thr Ala Asp Thr Ser Ser Asn  
85 90 95

Thr Ala Tyr Leu Gln Leu Ser Ser Leu Thr Ser Glu Asp Thr Ala Val  
100 105 110

Tyr Tyr Cys Ala Lys Pro Tyr Gly Asn Tyr Gly Tyr Tyr Tyr Ala Leu  
115 120 125

Asp Tyr Trp Gly Gln Gly Thr Ser Val Thr Val Ser Ser  
130 135 140

<210> 1326

<211> 90

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 heavy chain variable framework 1 (FW1)  
sequence

<400> 1326

gaggttcagc tgcagcagtc tggggcagac cttgtgaagc caggggcctc agtcaagttg 60

tcctgcacag cttctggcctt caacattaaa 90

<210> 1327

<211> 30

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 heavy chain variable framework 1 (FW1)  
sequence

<400> 1327

Glu Val Gln Leu Gln Gln Ser Gly Ala Asp Leu Val Lys Pro Gly Ala

1 5 10 15

Ser Val Lys Leu Ser Cys Thr Ala Ser Gly Phe Asn Ile Lys  
20 25 30

<210> 1328  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 45C11 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1328  
gacaccttta tgcac 15

<210> 1329  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 45C11 heavy chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1329

Asp Thr Phe Met His  
1 5

<210> 1330  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 45C11 heavy chain variable framework 2 (FW2)  
sequence

<400> 1330  
tgggtgaagc agaggcctga acagggcctg gaggaggattg ga 42

<210> 1331  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 45C11 heavy chain variable framework 2 (FW2)  
sequence

<400> 1331

Trp Val Lys Gln Arg Pro Glu Gln Gly Leu Glu Trp Ile Gly  
1 5 10

<210> 1332  
<211> 51  
<212> DNA  
<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1332

aggattgata ctgcgaatgg taataactaaa tatgacccga aattccaggg c

51

<210> 1333

<211> 17

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 heavy chain variable complementarity  
determining regions 2 (CDR2) sequence

<400> 1333

Arg Ile Asp Pro Ala Asn Gly Asn Thr Lys Tyr Asp Pro Lys Phe Gln

1

5

10

15

Gly

<210> 1334

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 heavy chain variable framework 3 (FW3)  
sequence

<400> 1334

aaggccacta taacagcaga cacatcctcc aacacagcct acctgcagct cagcagcctg

60

acatctgagg acactgccgt ctattactgt gctaaa

96

<210> 1335

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 heavy chain variable framework 3 (FW3)  
sequence

<400> 1335

Lys Ala Thr Ile Thr Ala Asp Thr Ser Ser Asn Thr Ala Tyr Leu Gln

1

5

10

15

Leu Ser Ser Leu Thr Ser Glu Asp Thr Ala Val Tyr Tyr Cys Ala Lys

20

25

30

<210> 1336

<211> 39

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1336  
ccgtatggta actacggcta ttactatgct ttggactac 39

<210> 1337  
<211> 13  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 45C11 heavy chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1337  
Pro Tyr Gly Asn Tyr Gly Tyr Tyr Tyr Ala Leu Asp Tyr  
1 5 10

<210> 1338  
<211> 33  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 45C11 heavy chain variable framework 4 (FW4)  
sequence

<400> 1338  
tggggtaag gaacctcagt caccgtctcc tca 33

<210> 1339  
<211> 11  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 45C11 heavy chain variable framework 4 (FW4)  
sequence

<400> 1339  
Trp Gly Gln Gly Thr Ser Val Thr Val Ser Ser  
1 5 10

<210> 1340  
<211> 381  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse antibody 45C11 Light chain: DNA sequence (381 bp) - Signal  
sequenceFR1CDR1FR2CDR2FR3CDR3FR4

<400> 1340  
atgaggttcc aggttcaggt tctggggctc cttctgctct ggatatcagg tgcccagtg 60  
gatgtccaga taaccagtc tccatcttat ctgctgcat ctctggaga aaccattact 120  
attaattgca gggcaagtaa gagcattagc aaatatttag cctggatatca agagaaacct 180  
gggaaaacta ataagcttct tatctactct ggatccactt tgcaatctgg aattccatca 240  
aggttcagtg gcagtggatc tggtagatg ttactctca ccatcagtag cctggagcct 300  
gaagattttg caatgtatta ctgtcaacag cataatgaat tcccgtggac gttcgggtgga 360



&lt;210&gt; 1341

&lt;211&gt; 127

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

<223> Mouse antibody 45C11 Light chain: Amino acid sequence (127 aa) -  
Signal peptideFR1CDR1FR2CDR2FR3CDR3FR4

&lt;400&gt; 1341

Met Arg Phe Gln Val Gln Val Leu Gly Leu Leu Leu Trp Ile Ser  
1 5 10 15Gly Ala Gln Cys Asp Val Gln Ile Thr Gln Ser Pro Ser Tyr Leu Ala  
20 25 30Ala Ser Pro Gly Glu Thr Ile Thr Ile Asn Cys Arg Ala Ser Lys Ser  
35 40 45Ile Ser Lys Tyr Leu Ala Trp Tyr Gln Glu Lys Pro Gly Lys Thr Asn  
50 55 60Lys Leu Leu Ile Tyr Ser Gly Ser Thr Leu Gln Ser Gly Ile Pro Ser  
65 70 75 80Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser  
85 90 95Ser Leu Glu Pro Glu Asp Phe Ala Met Tyr Tyr Cys Gln Gln His Asn  
100 105 110Glu Phe Pro Trp Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
115 120 125

&lt;210&gt; 1342

&lt;211&gt; 69

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

<223> Mouse antibody 45C11 light chain variable framework 1 (FW1)  
sequence

&lt;400&gt; 1342

gatgtccaga taaccagtc tccatcttat cttgctgcat ctcctggaga aaccattact 60

attaattgc 69

&lt;210&gt; 1343

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

<223> Mouse antibody 45C11 light chain variable framework 1 (FW1)  
sequence

<400> 1343

Asp Val Gln Ile Thr Gln Ser Pro Ser Tyr Leu Ala Ala Ser Pro Gly  
1 5 10 15

Glu Thr Ile Thr Ile Asn Cys  
20

<210> 1344

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1344

agggcaagta agagcattag caaatattta gcc

33

<210> 1345

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 light chain variable complementarity  
determining regions 1 (CDR1) sequence

<400> 1345

Arg Ala Ser Lys Ser Ile Ser Lys Tyr Leu Ala  
1 5 10

<210> 1346

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 light chain variable framework 2 (FW2)  
sequence

<400> 1346

tggtatcaag agaaacctgg gaaaactaat aagcttctta tctac

45

<210> 1347

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse antibody 45C11 light chain variable framework 2 (FW2)  
sequence

<400> 1347

Trp Tyr Gln Glu Lys Pro Gly Lys Thr Asn Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 1348

<211> 21

<212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse antibody 45C11 light chain variable complementarity  
 determining regions 2 (CDR2) sequence  
  
 <400> 1348  
 tctggatcca ctttgcaatc t 21

<210> 1349  
 <211> 7  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 45C11 light chain variable complementarity  
 determining regions 2 (CDR2) sequence

<400> 1349

Ser Gly Ser Thr Leu Gln Ser  
 1 5

<210> 1350  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 45C11 light chain variable framework 3 (FW3)  
 sequence

<400> 1350  
 ggaattccat caaggttcag tggcagtgga tctggtacag atttcactct caccatcagt 60  
 agcctggagc ctgaagattt tgcaatgtat tactgt 96

<210> 1351  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 45C11 light chain variable framework 3 (FW3)  
 sequence

<400> 1351

Gly Ile Pro Ser Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 1 5 10 15

Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe Ala Met Tyr Tyr Cys  
 20 25 30

<210> 1352  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse antibody 45C11 light chain variable complementarity  
 determining regions 3 (CDR3) sequence

<400> 1352  
caacagcata atgaattccc gtggacg 27

<210> 1353  
<211> 9  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 45C11 light chain variable complementarity  
determining regions 3 (CDR3) sequence

<400> 1353  
  
Gln Gln His Asn Glu Phe Pro Trp Thr  
1 5

<210> 1354  
<211> 30  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 45C11 light chain variable framework 4 (FW4)  
sequence

<400> 1354  
ttcgggtggag gcaccaagct ggaaatcaaa 30

<210> 1355  
<211> 10  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse antibody 45C11 light chain variable framework 4 (FW4)  
sequence

<400> 1355  
  
Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
1 5 10

<210> 1356  
<211> 90  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Mouse 5C6F3 heavy chain variable framework 1 (FW1) sequence

<400> 1356  
gaagtgatgc tgggtggagtc tggggggaggc ttagtgaagc ctggagggtc cctgaaactc 60  
tcctgtgcag cctctggatt cactttcagt 90

<210> 1357  
<211> 30  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse 5C6F3 heavy chain variable framework 1 (FW1) sequence

<400> 1357

Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 1358

<211> 15

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 heavy chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 1358

acctatgcc tgtct

15

<210> 1359

<211> 5

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 heavy chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 1359

Thr Tyr Ala Met Ser  
1 5

<210> 1360

<211> 42

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 heavy chain variable framework 2 (FW2) sequence

<400> 1360

tgggttcgcc agactccgga gaagaggctg gagggggtcg ca

42

<210> 1361

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 heavy chain variable framework 2 (FW2) sequence

<400> 1361

Trp Val Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val Ala  
1 5 10

<210> 1362

<211> 51

<212> DNA

<213> Artificial Sequence

<220>  
 <223> Mouse 5C6F3 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
 <400> 1362  
 gccattagta atggtggtgg ttacacctac tatccagaca gtctgaaggg g 51

<210> 1363  
 <211> 17  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse 5C6F3 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1363

Ala	Ile	Ser	Asn	Gly	Gly	Gly	Tyr	Thr	Tyr	Tyr	Pro	Asp	Ser	Leu	Lys
1				5					10					15	

Gly

<210> 1364  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse 5C6F3 heavy chain variable framework 3 (FW3) sequence

<400> 1364  
 cgattcacca tctccagaga caatgccaaag aacaccctgt acctgcaaat gagcagtctg 60  
 aggtctgagg acacggccac gtattactgt gcaaga 96

<210> 1365  
 <211> 32  
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<220>  
 <223> Mouse 5C6F3 heavy chain variable framework 3 (FW3) sequence

<400> 1365

Arg	Phe	Thr	Ile	Ser	Arg	Asp	Asn	Ala	Lys	Asn	Thr	Leu	Tyr	Leu	Gln
1				5					10					15	

Met	Ser	Ser	Leu	Arg	Ser	Glu	Asp	Thr	Ala	Thr	Tyr	Tyr	Cys	Ala	Arg
			20						25				30		

<210> 1366  
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<220>  
 <223> Mouse 5C6F3 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1366

cgttactatg atcactactt tgactac

27

<210> 1367

<211> 9

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1367

Arg Tyr Tyr Asp His Tyr Phe Asp Tyr

1 5

<210> 1368

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 heavy chain variable framework 4 (FW4) sequence

<400> 1368

tggggccaag gcaccgctct cacggtctcc tca

33

<210> 1369

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 heavy chain variable framework 4 (FW4) sequence

<400> 1369

Trp Gly Gln Gly Thr Ala Leu Thr Val Ser Ser

1 5 10

<210> 1370

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 light chain variable framework 1 (FR1) sequence

<400> 1370

gatgttttga tgacccaaac tccactctcc ctgcctgtca gtcttggaga tcaagcctcc

60

atctcttgc

69

<210> 1371

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 light chain variable framework 1 (FR1) sequence

<400> 1371

Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly

1 5 10 15

Asp Gln Ala Ser Ile Ser Cys  
20

<210> 1372

<211> 48

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1372

agatctagtc agaccattgt acatagtaat ggaaacacct atttagaa 48

<210> 1373

<211> 16

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1373

Arg Ser Ser Gln Thr Ile Val His Ser Asn Gly Asn Thr Tyr Leu Glu  
1 5 10 15

<210> 1374

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 light chain variable framework 2 (FR2) sequence

<400> 1374

tggtacctgc agaaaccagg ccagtctcca aagctcctga tctac 45

<210> 1375

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 light chain variable framework 2 (FR2) sequence

<400> 1375

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 1376

<211> 21

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 5C6F3 light chain variable complementarity determining regions 2 (CDR2) sequence



<400> 1376  
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<210> 1377  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 5C6F3 light chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1377

Lys Val Ser Asn Arg Phe Ser  
1 5

<210> 1378  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse 5C6F3 light chain variable framework 3 (FR3) sequence

<400> 1378  
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agggtggagg ctgaggatct gggagtttat tactgc 96

<210> 1379  
<211> 32  
<212> PRT  
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<220>  
<223> Mouse 5C6F3 light chain variable framework 3 (FR3) sequence

<400> 1379

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
20 25 30

<210> 1380  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse 5C6F3 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1380  
tttcaagatt cacatgttcc tctcacg 27

<210> 1381  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 5C6F3 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1381

Phe Gln Asp Ser His Val Pro Leu Thr  
1 5

<210> 1382  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse 5C6F3 light chain variable framework 4 (FR4) sequence

<400> 1382  
ttcgggtgctg ggaccaagct ggagctgaaa 30

<210> 1383  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 5C6F3 light chain variable framework 4 (FR4) sequence

<400> 1383  
Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys  
1 5 10

<210> 1384  
<211> 735  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> mu5C6F3 scFv sequence

<400> 1384  
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tcctgtgcag cctctggatt cactttcagt acctatgcc tgccttggtg tcgccagact 120  
ccggagaaga ggctggagtg ggtcgcagcc attagtaatg gtgggtggtta cacctactat 180  
ccagacagtc tgaaggggagc attcaccatc tccagagaca atgccaagaa caccctgtac 240  
ctgcaaatga gcagtctgag gtctgaggac acggccacgt attactgtgc aagacgttac 300  
tatgatcact actttgacta ctggggccaa ggcaccgctc tcacgggtctc ctcagggtggc 360  
ggaggatctg gcggaggtgg aagcggcgga ggcggatccg atgttttgat gacccaaact 420  
ccactctccc tgcctgtcag tcttgagat caagcctcca tctcttcag atctagtcag 480  
accattgtac atagtaatgg aaacacctat ttagaatggt acctgcagaa accaggccag 540  
tctccaaagc tcctgatcta caaagtttcc aaccgatttt ctgggggtccc agacaggttc 600  
agtggcagtg gatcaggagc agatttcaca ctcaagatca gcagggtgga ggctgaggat 660  
ctgggagttt attactgctt tcaagattca catgttcctc tcacgttcgg tgctgggacc 720

&lt;210&gt; 1385

&lt;211&gt; 245

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; mu5C6F3 scFv sequence

&lt;400&gt; 1385

Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Thr Tyr  
20 25 30

Ala Met Ser Trp Val Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val  
35 40 45

Ala Ala Ile Ser Asn Gly Gly Gly Tyr Thr Tyr Tyr Pro Asp Ser Leu  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr  
65 70 75 80

Leu Gln Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Thr Tyr Tyr Cys  
85 90 95

Ala Arg Arg Tyr Tyr Asp His Tyr Phe Asp Tyr Trp Gly Gln Gly Thr  
100 105 110

Ala Leu Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser  
115 120 125

Gly Gly Gly Gly Ser Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu  
130 135 140

Pro Val Ser Leu Gly Asp Gln Ala Ser Ile Ser Cys Arg Ser Ser Gln  
145 150 155 160

Thr Ile Val His Ser Asn Gly Asn Thr Tyr Leu Glu Trp Tyr Leu Gln  
165 170 175

Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr Lys Val Ser Asn Arg  
180 185 190

Phe Ser Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp  
195 200 205

Phe Thr Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr  
210 215 220

Tyr Cys Phe Gln Asp Ser His Val Pro Leu Thr Phe Gly Ala Gly Thr  
225 230 235 240

Lys Leu Glu Leu Lys  
245

<210> 1386  
<211> 90  
<212> DNA  
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<220>  
<223> Mouse 3C2B1 heavy chain variable framework 1 (CDR1) sequence

<400> 1386  
gaagtgatgc tgggtggagtc tggggggaggc ttagtgaagc ctggagggtc cctgaaactc 60  
tcctgtgcag cctctggaat cactttcagt 90

<210> 1387  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 3C2B1 heavy chain variable framework 1 (CDR1) sequence

<400> 1387

Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Ile Thr Phe Ser  
20 25 30

<210> 1388  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse 3C2B1 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1388  
acctatacca tgtcg 15

<210> 1389  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 3C2B1 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1389

Thr Tyr Thr Met Ser  
1 5

<210> 1390  
 <211> 42  
 <212> DNA  
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 <220>  
 <223> Mouse 3C2B1 heavy chain variable framework 2 (CDR2) sequence  
  
 <400> 1390  
 tgggttcgcc agactccgga gaagaggctg gagtgggtcg ca 42

<210> 1391  
 <211> 14  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 3C2B1 heavy chain variable framework 2 (CDR2) sequence  
  
 <400> 1391  
  
 Trp Val Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val Ala  
 1 5 10

<210> 1392  
 <211> 51  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 3C2B1 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 1392  
 accattagta ctggtggtga taaaacctac tattcagaca gtgtgaaggg t 51

<210> 1393  
 <211> 17  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 3C2B1 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 1393  
  
 Thr Ile Ser Thr Gly Gly Asp Lys Thr Tyr Tyr Ser Asp Ser Val Lys  
 1 5 10 15

Gly

<210> 1394  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 3C2B1 heavy chain variable framework 3 (CDR3) sequence  
  
 <400> 1394  
 cgattcacca tctccagaga caatgccaaag aacaacctgt acctccaaat gagcagtctg 60

aggctctgagg acacggcctt gtattactgt gcaagg

96

<210> 1395

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 3C2B1 heavy chain variable framework 3 (CDR3) sequence

<400> 1395

Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Asn Leu Tyr Leu Gln  
1 5 10 15

Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Leu Tyr Tyr Cys Ala Arg  
20 25 30

<210> 1396

<211> 36

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 3C2B1 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1396

ggaaccacgg ctatgtatta ctatgctatg gactac

36

<210> 1397

<211> 12

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 3C2B1 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1397

Gly Thr Thr Ala Met Tyr Tyr Tyr Ala Met Asp Tyr  
1 5 10

<210> 1398

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 3C2B1 heavy chain variable framework 4 (CDR4) sequence

<400> 1398

tgggggtcaag gaacctcagt caccgtctcc tca

33

<210> 1399

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 3C2B1 heavy chain variable framework 4 (CDR4) sequence

<400> 1399

Trp Gly Gln Gly Thr Ser Val Thr Val Ser Ser  
1 5 10

<210> 1400

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 3C2B1 light chain variable framework 1 (FW1) sequence

<400> 1400

gacattgtgc tgacacagtc tcctgcttcc ttagctgtat ctctggggca gagggccacc 60

atctcatgc 69

<210> 1401

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 3C2B1 light chain variable framework 1 (FW1) sequence

<400> 1401

Asp Ile Val Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Leu Gly  
1 5 10 15

Gln Arg Ala Thr Ile Ser Cys  
20

<210> 1402

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 3C2B1 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1402

agggccagca aaagtatcag tacatctgac tataattata ttcac 45

<210> 1403

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 3C2B1 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1403

Arg Ala Ser Lys Ser Ile Ser Thr Ser Asp Tyr Asn Tyr Ile His  
1 5 10 15

<210> 1404

<211> 45

<212> DNA

<213> Artificial Sequence  
 <220>  
 <223> Mouse 3C2B1 light chain variable framework 2 (FW2) sequence  
 <400> 1404  
 tggtagcaaac agaaaccagg acagccaccc aaactcctca tctat 45  
 <210> 1405  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence  
 <220>  
 <223> Mouse 3C2B1 light chain variable framework 2 (FW2) sequence  
 <400> 1405  
 Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr  
 1 5 10 15  
 <210> 1406  
 <211> 21  
 <212> DNA  
 <213> Artificial Sequence  
 <220>  
 <223> Mouse 3C2B1 light chain variable complementarity determining  
 regions 2 (CDR2) sequence  
 <400> 1406  
 cttgcatcca acctagaatc t 21  
 <210> 1407  
 <211> 7  
 <212> PRT  
 <213> Artificial Sequence  
 <220>  
 <223> Mouse 3C2B1 light chain variable complementarity determining  
 regions 2 (CDR2) sequence  
 <400> 1407  
 Leu Ala Ser Asn Leu Glu Ser  
 1 5  
 <210> 1408  
 <211> 95  
 <212> DNA  
 <213> Artificial Sequence  
 <220>  
 <223> Mouse 3C2B1 light chain variable framework 3 (FW3) sequence  
 <400> 1408  
 gggtagcctgc caggttcagt ggcagtaggt ctgggacaga cttcacctc aacatccatc 60  
 ctgtggagga agaagatgct gcaacctatt actgt 95  
 <210> 1409  
 <211> 32  
 <212> PRT  
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<220>  
<223> Mouse 3C2B1 light chain variable framework 3 (FW3) sequence

<400> 1409

Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Asn Ile His Pro Val Glu Glu Glu Asp Ala Ala Thr Tyr Tyr Cys  
20 25 30

<210> 1410  
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<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse 3C2B1 light chain variable complementarity determining  
regions 3 (CDR3) sequence

<400> 1410  
cagcacagta gggagcttcc tctcacg 27

<210> 1411  
<211> 9  
<212> PRT  
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<220>  
<223> Mouse 3C2B1 light chain variable complementarity determining  
regions 3 (CDR3) sequence

<400> 1411

Gln His Ser Arg Glu Leu Pro Leu Thr  
1 5

<210> 1412  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse 3C2B1 light chain variable framework 4 (FW4) sequence

<400> 1412  
ttcggtgctg ggaccaagct ggagctgaaa 30

<210> 1413  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 3C2B1 light chain variable framework 4 (FW4) sequence

<400> 1413

Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys  
1 5 10

<210> 1414

<211> 90  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse B12 heavy chain variable framework 1 (FW1) sequence

<400> 1414  
 caggcgagc tgaaggagtc aggacctggc ctggtggcgc cctcacagag cctgtccatc 60  
 acttgcaactg tctctgggtt ttcattaacc 90

<210> 1415  
 <211> 30  
 <212> PRT  
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<220>  
 <223> Mouse B12 heavy chain variable framework 1 (FW1) sequence

<400> 1415  
 Gln Ala Gln Leu Lys Glu Ser Gly Pro Gly Leu Val Ala Pro Ser Gln  
 1 5 10 15  
 Ser Leu Ser Ile Thr Cys Thr Val Ser Gly Phe Ser Leu Thr  
 20 25 30

<210> 1416  
 <211> 15  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse B12 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1416  
 agctatggtg tacac 15

<210> 1417  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse B12 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1417  
 Ser Tyr Gly Val His  
 1 5

<210> 1418  
 <211> 42  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse B12 heavy chain variable framework 2 (FW2) sequence

<400> 1418  
 tgggttcgcc agcctccagg aaagggtctg gagggtggctgg ga 42

<210> 1419  
<211> 14  
<212> PRT  
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<220>  
<223> Mouse B12 heavy chain variable framework 2 (FW2) sequence

<400> 1419

Trp Val Arg Gln Pro Pro Gly Lys Gly Leu Glu Trp Leu Gly  
1 5 10

<210> 1420  
<211> 54  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B12 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1420  
gtaatatggc ctggtggaag cacaaattat aattcgactc tcatgtccag aatg 54

<210> 1421  
<211> 18  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B12 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1421

Val Ile Trp Pro Gly Gly Ser Thr Asn Tyr Asn Ser Thr Leu Met Ser  
1 5 10 15

Arg Met

<210> 1422  
<211> 90  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B12 heavy chain variable framework 3 (FW3) sequence

<400> 1422  
cggatcatca aagacaactc caagagccaa gttttcttaa aaatgaacag tctgcaaatt 60

gatgacacag ccatgtacta ctgtgccaga 90

<210> 1423  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B12 heavy chain variable framework 3 (FW3) sequence

<400> 1423

Arg Ile Ile Lys Asp Asn Ser Lys Ser Gln Val Phe Leu Lys Met Asn  
1 5 10 15

Ser Leu Gln Ile Asp Asp Thr Ala Met Tyr Tyr Cys Ala Arg  
20 25 30

<210> 1424

<211> 36

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B12 heavy chain variable complementarity determining  
regions 3 (CDR3) sequence

<400> 1424

gacgcggacac ctcgggtggg ggcctggttt gcttac

36

<210> 1425

<211> 12

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B12 heavy chain variable complementarity determining  
regions 3 (CDR3) sequence

<400> 1425

Asp Arg Thr Pro Arg Val Gly Ala Trp Phe Ala Tyr  
1 5 10

<210> 1426

<211> 34

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B12 heavy chain variable framework 4 (FW4) sequence

<400> 1426

tggggccaag ggactctggt cactgtctct gcag

34

<210> 1427

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B12 heavy chain variable framework 4 (FW4) sequence

<400> 1427

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
1 5 10

<210> 1428

<211> 69

<212> DNA

<213> Artificial Sequence

<220>  
 <223> Mouse B12 light chain variable framework 1 (FR1) sequence  
  
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 atatcctgc 69  
  
 <210> 1429  
 <211> 23  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B12 light chain variable framework 1 (FR1) sequence  
  
 <400> 1429  
  
 Ile Ile Val Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Leu Gly  
 1 5 10 15  
  
 Gln Arg Ala Thr Ile Ser Cys  
 20  
  
 <210> 1430  
 <211> 45  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B12 light chain variable complementarity determining  
 regions 1 (CDR1) sequence  
  
 <400> 1430  
 agagccagtg agagtgttgc tacttatggc aataatttta tgcag 45  
  
 <210> 1431  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B12 light chain variable complementarity determining  
 regions 1 (CDR1) sequence  
  
 <400> 1431  
  
 Arg Ala Ser Glu Ser Val Ala Thr Tyr Gly Asn Asn Phe Met Gln  
 1 5 10 15  
  
 <210> 1432  
 <211> 90  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B2 heavy chain variable framework 1 (FW1) sequence  
  
 <400> 1432  
 gaagtgggtgc tgggtggagtc tgggggaggc ttagtggagc ctggagggtc cctgaaactc 60  
 tcctgtgtag cctctggatt cgctttcagt 90

<210> 1433  
 <211> 30  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B2 heavy chain variable framework 1 (FW1) sequence  
  
 <400> 1433

Glu Val Val Leu Val Glu Ser Gly Gly Gly Leu Val Glu Pro Gly Gly  
 1 5 10 15

Ser Leu Lys Leu Ser Cys Val Ala Ser Gly Phe Ala Phe Ser  
 20 25 30

<210> 1434  
 <211> 15  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B2 heavy chain variable complementarity determining regions  
 1 (CDR1) sequence

<400> 1434  
 acctttgcc tgtct 15

<210> 1435  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B2 heavy chain variable complementarity determining regions  
 1 (CDR1) sequence

<400> 1435  
  
 Thr Phe Ala Met Ser  
 1 5

<210> 1436  
 <211> 42  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse B2 heavy chain variable framework 2 (FW2) sequence

<400> 1436  
 tggattcgcc agactccgga gaagaggctg gagtgggtcg ca 42

<210> 1437  
 <211> 14  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse B2 heavy chain variable framework 2 (FW2) sequence

<400> 1437

Trp Ile Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val Ala  
1 5 10

<210> 1438

<211> 51

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B2 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1438

gccattagta atggtggtgg ttacacctac tatccagaca ctctgaagg g 51

<210> 1439

<211> 17

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B2 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1439

Ala Ile Ser Asn Gly Gly Gly Tyr Thr Tyr Tyr Pro Asp Thr Leu Lys  
1 5 10 15

Gly

<210> 1440

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B2 heavy chain variable framework 3 (FW3) sequence

<400> 1440

cgattctcca tctccagaga caatgccaa g aataccctgt acctgcaa at gagtagtctg 60

aggtctgagg acacggccgt gtattactgt gcaaga 96

<210> 1441

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B2 heavy chain variable framework 3 (FW3) sequence

<400> 1441

Arg Phe Ser Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln  
1 5 10 15

Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Val Tyr Tyr Cys Ala Arg  
20 25 30

<210> 1442

<211> 27  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B2 heavy chain variable complementarity determining regions  
 3 (CDR3) sequence  
  
 <400> 1442  
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<210> 1443  
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 <223> Mouse B2 heavy chain variable complementarity determining regions  
 3 (CDR3) sequence  
  
 <400> 1443  
  
 Arg Tyr Tyr Asp Leu Tyr Phe Asp Leu  
 1 5

<210> 1444  
 <211> 33  
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 <213> Artificial Sequence  
  
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 <223> Mouse B2 heavy chain variable framework 4 (FW4) sequence  
  
 <400> 1444  
 tggggccgag gcacctctct catagtctcc tca 33

<210> 1445  
 <211> 11  
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 <213> Artificial Sequence  
  
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 <223> Mouse B2 heavy chain variable framework 4 (FW4) sequence  
  
 <400> 1445  
  
 Trp Gly Arg Gly Thr Ser Leu Ile Val Ser Ser  
 1 5 10

<210> 1446  
 <211> 69  
 <212> DNA  
 <213> Artificial Sequence  
  
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 <223> Mouse B2 light chain variable framework 1 (FR1) sequence  
  
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 atttcttgc 69

<210> 1447  
 <211> 23



<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B2 light chain variable framework 1 (FR1) sequence

<400> 1447

Asp Ile Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
1 5 10 15

Asp Gln Ala Ser Ile Ser Cys  
20

<210> 1448  
<211> 48  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B2 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1448  
agatcttagtc agaacattgt acatagtaat ggaaacacct atttagaa 48

<210> 1449  
<211> 16  
<212> PRT  
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<220>  
<223> Mouse B2 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1449

Arg Ser Ser Gln Asn Ile Val His Ser Asn Gly Asn Thr Tyr Leu Glu  
1 5 10 15

<210> 1450  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B2 light chain variable framework 2 (FR2) sequence

<400> 1450  
tggtacctgc agaaaccagg ccagtcctcca aagctcctga tctac 45

<210> 1451  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B2 light chain variable framework 2 (FR2) sequence

<400> 1451

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 1452  
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 <213> Artificial Sequence  
  
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 <223> Mouse B2 light chain variable complementarity determining regions  
 2 (CDR2) sequence  
  
 <400> 1452  
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 <210> 1453  
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 <212> PRT  
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 <220>  
 <223> Mouse B2 light chain variable complementarity determining regions  
 2 (CDR2) sequence  
  
 <400> 1453  
  
 Lys Val Ser Asn Arg Phe Ser  
 1 5  
  
 <210> 1454  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B2 light chain variable framework 3 (FR3) sequence  
  
 <400> 1454  
 ggggtccccg acaggtttcag tggtagtggg tcagggacag atttcacact caagatcagc 60  
 agagtggagg ctgaggatct gggagtttat tactgc 96  
  
 <210> 1455  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B2 light chain variable framework 3 (FR3) sequence  
  
 <400> 1455  
  
 Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 1 5 10 15  
  
 Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
 20 25 30  
  
 <210> 1456  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B2 light chain variable complementarity determining regions  
 3 (CDR3) sequence

<400> 1456  
tttcaagatt cacatgttcc tctcacg 27

<210> 1457  
<211> 9  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse B2 light chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 1457  
  
Phe Gln Asp Ser His Val Pro Leu Thr  
1 5

<210> 1458  
<211> 30  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Mouse B2 light chain variable framework 4 (FR4) sequence

<400> 1458  
ttcgggtgctg ggaccaggct ggagctgaaa 30

<210> 1459  
<211> 10  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse B2 light chain variable framework 4 (FR4) sequence  
  
<400> 1459

Phe Gly Ala Gly Thr Arg Leu Glu Leu Lys  
1 5 10

<210> 1460  
<211> 90  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Mouse B7 heavy chain variable framework 1 (FW1) sequence  
  
<400> 1460  
gaggtgcagg tgggtggagtc tgggggagac ttagtgaagc ctggagggtc cctgaaactc 60  
  
tcctgtgcag cctctggatt cactttcagt 90

<210> 1461  
<211> 30  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse B7 heavy chain variable framework 1 (FW1) sequence  
  
<400> 1461

Glu Val Gln Val Val Glu Ser Gly Gly Asp Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 1462  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B7 heavy chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1462  
agatatggca tgtct 15

<210> 1463  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B7 heavy chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1463

Arg Tyr Gly Met Ser  
1 5

<210> 1464  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B7 heavy chain variable framework 2 (FW2) sequence

<400> 1464  
tgggttcgcc agactccaga caagaggctg gagtgggtcg ca 42

<210> 1465  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B7 heavy chain variable framework 2 (FW2) sequence

<400> 1465

Trp Val Arg Gln Thr Pro Asp Lys Arg Leu Glu Trp Val Ala  
1 5 10

<210> 1466  
<211> 51  
<212> DNA  
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<220>

<223> Mouse B7 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1466  
accattagta gtggtggtac ttacatctac tatccagaca gtgtgaaggg g 51

<210> 1467  
<211> 17  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B7 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1467  
Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys  
1 5 10 15

Gly

<210> 1468  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B7 heavy chain variable framework 3 (FW3) sequence

<400> 1468  
cgattcacca tctccagaga caatgccaaag aacaccctgt acctgcaaat gagcagtctg 60  
aagtctgagg acacagccat gtattactgt gcaagg 96

<210> 1469  
<211> 32  
<212> PRT  
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<220>  
<223> Mouse B7 heavy chain variable framework 3 (FW3) sequence

<400> 1469  
Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln  
1 5 10 15

Met Ser Ser Leu Lys Ser Glu Asp Thr Ala Met Tyr Tyr Cys Ala Arg  
20 25 30

<210> 1470  
<211> 39  
<212> DNA  
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<220>  
<223> Mouse B7 heavy chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 1470  
gataactacg gtagtagcta cgactatgct atggactac 39

<210> 1471  
<211> 13  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B7 heavy chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 1471

Asp Asn Tyr Gly Ser Ser Tyr Asp Tyr Ala Met Asp Tyr  
1 5 10

<210> 1472  
<211> 33  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B7 heavy chain variable framework 4 (FW4) sequence

<400> 1472  
tgggggtcaag gaacctcagt caccgtctcc tca 33

<210> 1473  
<211> 11  
<212> PRT  
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<220>  
<223> Mouse B7 heavy chain variable framework 4 (FW4) sequence

<400> 1473

Trp Gly Gln Gly Thr Ser Val Thr Val Ser Ser  
1 5 10

<210> 1474  
<211> 69  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B7 light chain variable framework 1 (FR1) sequence

<400> 1474  
gatgttttga tgacccaaac tccactctcc ctgcctgtca gtcttggaga tcaagcctcc 60  
atctcttgc 69

<210> 1475  
<211> 23  
<212> PRT  
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<220>  
<223> Mouse B7 light chain variable framework 1 (FR1) sequence

<400> 1475

Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
1 5 10 15

Asp Gln Ala Ser Ile Ser Cys  
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<210> 1476

<211> 48

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B7 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1476

agatctagtc agaccattgt acatagtaat ggaaacacct atttagaa 48

<210> 1477

<211> 16

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B7 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1477

Arg Ser Ser Gln Thr Ile Val His Ser Asn Gly Asn Thr Tyr Leu Glu  
1 5 10 15

<210> 1478

<211> 45

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B7 light chain variable framework 2 (FR2) sequence

<400> 1478

tggtagctgc aaaaaccagg ccagttctcca aagctcctga tctac 45

<210> 1479

<211> 15

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B7 light chain variable framework 2 (FR2) sequence

<400> 1479

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 1480

<211> 21

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B7 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1480  
aaagtttcca accgattttc t 21

<210> 1481  
<211> 7  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse B7 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1481  
  
Lys Val Ser Asn Arg Phe Ser  
1 5

<210> 1482  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B7 light chain variable framework 3 (FR3) sequence

<400> 1482  
gggggtcccag acaggttcag tggcagtgga tcagggacag atttcacact caagatcagc 60  
  
agggtggagg ctgaggatct gggagtttat tactgc 96

<210> 1483  
<211> 32  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> Mouse B7 light chain variable framework 3 (FR3) sequence  
  
<400> 1483

Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
20 25 30

<210> 1484  
<211> 27  
<212> DNA  
<213> Artificial Sequence  
  
<220>  
<223> Mouse B7 light chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 1484  
tttcaagatt cacatgttcc tctcacg 27

<210> 1485  
<211> 9  
<212> PRT  
<213> Artificial Sequence



<220>  
 <223> Mouse B7 light chain variable complementarity determining regions  
 3 (CDR3) sequence  
  
 <400> 1485  
  
 Phe Gln Asp Ser His Val Pro Leu Thr  
 1 5  
  
 <210> 1486  
 <211> 30  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B7 light chain variable framework 4 (FR4) sequence  
  
 <400> 1486  
 ttcggtgctg ggaccaagct ggagctgaaa 30  
  
 <210> 1487  
 <211> 10  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B7 light chain variable framework 4 (FR4) sequence  
  
 <400> 1487  
  
 Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys  
 1 5 10  
  
 <210> 1488  
 <211> 90  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 8C7F3 heavy chain variable framework 1 (FW1) sequence  
  
 <400> 1488  
 gaagtgatgc tggaggagtc tgggggaggc ttagtgaagc ctggagggtc cctgaaactc 60  
 tcctgtgcag cctctggatt cactttcagt 90  
  
 <210> 1489  
 <211> 30  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 8C7F3 heavy chain variable framework 1 (FW1) sequence  
  
 <400> 1489  
  
 Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
 1 5 10 15  
  
 Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
 20 25 30  
  
 <210> 1490

<211> 15  
 <212> DNA  
 <213> Artificial Sequence  
  
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 <223> Mouse 8C7F3 heavy chain variable complementarity determining regions 1 (CDR1) sequence  
  
 <400> 1490  
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15

<210> 1491  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 8C7F3 heavy chain variable complementarity determining regions 1 (CDR1) sequence  
  
 <400> 1491  
  
 Thr Tyr Ala Met Ser  
 1 5  
  
 <210> 1492  
 <211> 42  
 <212> DNA  
 <213> Artificial Sequence  
  
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 <223> Mouse 8C7F3 heavy chain variable framework 2 (FW2) sequence  
  
 <400> 1492  
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42

<210> 1493  
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 <212> PRT  
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 <223> Mouse 8C7F3 heavy chain variable framework 2 (FW2) sequence  
  
 <400> 1493  
  
 Trp Val Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val Ala  
 1 5 10  
  
 <210> 1494  
 <211> 51  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 8C7F3 heavy chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 1494  
 gccattagta atggtggtgg ttacacctac tatccagaca gtctgaagg g

51

<210> 1495  
 <211> 17  
 <212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1495

Ala Ile Ser Asn Gly Gly Gly Tyr Thr Tyr Tyr Pro Asp Ser Leu Lys  
1 5 10 15

Gly

<210> 1496

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 heavy chain variable framework 3 (FW3) sequence

<400> 1496

cgattcacca tctccagaga caatgccaaag aacaccctgt acctgcaaat gagcagtctg 60

aggctctgagg acacggccac gtattactgt gcaaga 96

<210> 1497

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 heavy chain variable framework 3 (FW3) sequence

<400> 1497

Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln  
1 5 10 15

Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Thr Tyr Tyr Cys Ala Arg  
20 25 30

<210> 1498

<211> 27

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1498

cgttactatg atcactactt tgactac 27

<210> 1499

<211> 9

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1499

Arg Tyr Tyr Asp His Tyr Phe Asp Tyr  
1 5

<210> 1500

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 heavy chain variable framework 4 (FW4) sequence

<400> 1500

tggggccaag gcaccgctct cacggtctcc tca 33

<210> 1501

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 heavy chain variable framework 4 (FW4) sequence

<400> 1501

Trp Gly Gln Gly Thr Ala Leu Thr Val Ser Ser  
1 5 10

<210> 1502

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 light chain variable framework 1 (FR1) sequence

<400> 1502

atcattgtgc tgacccaatc tccagcttct ttggctgtgt ctctagggca gagggccacc 60

atatacctgc 69

<210> 1503

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 light chain variable framework 1 (FR1) sequence

<400> 1503

Ile Ile Val Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Leu Gly  
1 5 10 15

Gln Arg Ala Thr Ile Ser Cys  
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<210> 1504

<211> 45

<212> DNA

<213> Artificial Sequence

<220>  
 <223> Mouse 8C7F3 light chain variable complementarity determining regions 1 (CDR1) sequence  
  
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 <210> 1505  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 8C7F3 light chain variable complementarity determining regions 1 (CDR1) sequence  
  
 <400> 1505  
  
 Arg Ala Ser Glu Ser Val Ala Thr Tyr Gly Asn Asn Phe Met Gln  
 1 5 10 15  
  
 <210> 1506  
 <211> 45  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 8C7F3 light chain variable framework 2 (FR2) sequence  
  
 <400> 1506  
 tggatcagc agaaaccagg acagccaccc aaactcctca tctat 45  
  
 <210> 1507  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 8C7F3 light chain variable framework 2 (FR2) sequence  
  
 <400> 1507  
  
 Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr  
 1 5 10 15  
  
 <210> 1508  
 <211> 21  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 8C7F3 light chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 1508  
 cttgcatcca ccctagattc t 21  
  
 <210> 1509  
 <211> 7  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>

<223> Mouse 8C7F3 light chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1509

Leu Ala Ser Thr Leu Asp Ser  
1 5

<210> 1510

<211> 96

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 light chain variable framework 3 (FR3) sequence

<400> 1510

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cctgtggagg ctgatgatgc tgcaacctat tactgt 96

<210> 1511

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 light chain variable framework 3 (FR3) sequence

<400> 1511

Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Arg Thr Asp Phe Thr  
1 5 10 15

Leu Thr Ile Asp Pro Val Glu Ala Asp Asp Ala Ala Thr Tyr Tyr Cys  
20 25 30

<210> 1512

<211> 27

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1512

cagcaaaata atgaggatcc tccgacg 27

<210> 1513

<211> 9

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse 8C7F3 light chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1513

Gln Gln Asn Asn Glu Asp Pro Pro Thr  
1 5

<210> 1514  
 <211> 30  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse 8C7F3 light chain variable framework 4 (FR4) sequence  
  
 <400> 1514  
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<210> 1515  
 <211> 10  
 <212> PRT  
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 <220>  
 <223> Mouse 8C7F3 light chain variable framework 4 (FR4) sequence  
  
 <400> 1515  
  
 Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
 1 5 10

<210> 1516  
 <211> 90  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse H11 heavy chain variable framework 1 (FW1) sequence  
  
 <400> 1516  
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 tcctgtgtag cctctggatt cgcttttagt 90

<210> 1517  
 <211> 30  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse H11 heavy chain variable framework 1 (FW1) sequence  
  
 <400> 1517  
  
 Glu Val Val Leu Val Glu Ser Gly Gly Gly Leu Val Glu Pro Gly Gly  
 1 5 10 15  
  
 Ser Leu Lys Leu Ser Cys Val Ala Ser Gly Phe Ala Phe Ser  
 20 25 30

<210> 1518  
 <211> 15  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse H11 heavy chain variable complementarity determining  
 regions 1 (CDR1) sequence  
  
 <400> 1518  
 acctttgcc tgcct 15

<210> 1519  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse H11 heavy chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1519

Thr Phe Ala Met Ser  
1 5

<210> 1520  
<211> 42  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse H11 heavy chain variable framework 2 (FW2) sequence

<400> 1520  
tggattcgcc agactccgga gaagaggctg gagggggtcg ca 42

<210> 1521  
<211> 14  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse H11 heavy chain variable framework 2 (FW2) sequence

<400> 1521

Trp Ile Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val Ala  
1 5 10

<210> 1522  
<211> 51  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse H11 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1522  
gccattagta atggtggtgg ttacacttac tatccagaca ctctgaagg g 51

<210> 1523  
<211> 17  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse H11 heavy chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1523

Ala Ile Ser Asn Gly Gly Gly Tyr Thr Tyr Tyr Pro Asp Thr Leu Lys  
1 5 10 15



Gly

<210> 1524  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse H11 heavy chain variable framework 3 (FW3) sequence

<400> 1524  
cgattcacca tctccagaga caatgccaaag aataccctgt acctgcaaat gagtagtctg 60  
aggctctgagg acacggccgt gtattactgt gcaaga 96

<210> 1525  
<211> 32  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse H11 heavy chain variable framework 3 (FW3) sequence

<400> 1525

Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln  
1 5 10 15

Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Val Tyr Tyr Cys Ala Arg  
20 25 30

<210> 1526  
<211> 27  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse H11 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1526  
cgctactatg atctctactt tgactta 27

<210> 1527  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse H11 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1527

Arg Tyr Tyr Asp Leu Tyr Phe Asp Leu  
1 5

<210> 1528  
<211> 33  
<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse H11 heavy chain variable framework 4 (FW4) sequence

<400> 1528

tggggccaag gcacctctct catagtctcc tca

33

<210> 1529

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse H11 heavy chain variable framework 4 (FW4) sequence

<400> 1529

Trp Gly Gln Gly Thr Ser Leu Ile Val Ser Ser  
1 5 10

<210> 1530

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse H11 light chain variable framework 1 (FR1) sequence

<400> 1530

gatattctga tgacccaaac tccactctcc ctgcctgtca gtcttggaga tcaagcctcc

60

atttcttgc

69

<210> 1531

<211> 23

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse H11 light chain variable framework 1 (FR1) sequence

<400> 1531

Asp Ile Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly  
1 5 10 15

Asp Gln Ala Ser Ile Ser Cys  
20

<210> 1532

<211> 48

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse H11 light chain variable complementarity determining  
regions 1 (CDR1) sequence

<400> 1532

agatctagtc agaacattgt acatagtaat ggaaacacct atttagaa

48

<210> 1533

<211> 16  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse H11 light chain variable complementarity determining regions 1 (CDR1) sequence

<400> 1533

Arg Ser Ser Gln Asn Ile Val His Ser Asn Gly Asn Thr Tyr Leu Glu  
1 5 10 15

<210> 1534  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse H11 light chain variable framework 2 (FR2) sequence

<400> 1534  
tggtacctgc agaaaccagg ccagtcctcca aagctcctga tctac 45

<210> 1535  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse H11 light chain variable framework 2 (FR2) sequence

<400> 1535

Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr  
1 5 10 15

<210> 1536  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse H11 light chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1536  
aaagtttcca accgattttc t 21

<210> 1537  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse H11 light chain variable complementarity determining regions 2 (CDR2) sequence

<400> 1537

Lys Val Ser Asn Arg Phe Ser  
1 5

<210> 1538

<211> 96  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse H11 light chain variable framework 3 (FR3) sequence  
  
 <400> 1538  
 ggggtccccg acaggttcag tggtagtggg tcaggacag atttcacact caagatcagc 60  
  
 agagtggagg ctgaggatct gggagtttat tactgc 96

<210> 1539  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse H11 light chain variable framework 3 (FR3) sequence  
  
 <400> 1539  
  
 Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr  
 1 5 10 15

Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys  
 20 25 30

<210> 1540  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse H11 light chain variable complementarity determining  
 regions 3 (CDR3) sequence  
  
 <400> 1540  
 tttcaagatt cacatgttcc tctcacg 27

<210> 1541  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse H11 light chain variable complementarity determining  
 regions 3 (CDR3) sequence  
  
 <400> 1541

Phe Gln Asp Ser His Val Pro Leu Thr  
 1 5

<210> 1542  
 <211> 30  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse H11 light chain variable framework 4 (FR4) sequence  
  
 <400> 1542  
 ttcggtgctg ggaccaggct ggagctgaaa 30

<210> 1543  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse H11 light chain variable framework 4 (FR4) sequence

<400> 1543

Phe Gly Ala Gly Thr Arg Leu Glu Leu Lys  
1 5 10

<210> 1544  
<211> 90  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B9 heavy chain variable framework 1 (FW1) sequence

<400> 1544  
gaggtgcagg tgggtggagtc tgggggagac ttagtgaagc ctggagggtc cctgaaactc 60  
tcctgtgcag cctctggatt cactttcagt 90

<210> 1545  
<211> 30  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B9 heavy chain variable framework 1 (FW1) sequence

<400> 1545

Glu Val Gln Val Val Glu Ser Gly Gly Asp Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser  
20 25 30

<210> 1546  
<211> 15  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B9 heavy chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1546  
agatatggca tgtct 15

<210> 1547  
<211> 5  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B9 heavy chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1547

Arg Tyr Gly Met Ser  
1 5

<210> 1548

<211> 42

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B9 heavy chain variable framework 2 (FW2) sequence

<400> 1548

tgggttcgcc agactccaga caagaggctg gagtgggtcg ca 42

<210> 1549

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B9 heavy chain variable framework 2 (FW2) sequence

<400> 1549

Trp Val Arg Gln Thr Pro Asp Lys Arg Leu Glu Trp Val Ala  
1 5 10

<210> 1550

<211> 51

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B9 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1550

accattagta gtggtgttac ttacatctac tatccagaca gtgtgaaggg g 51

<210> 1551

<211> 17

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B9 heavy chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1551

Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr Tyr Pro Asp Ser Val Lys  
1 5 10 15

Gly

<210> 1552

<211> 93

<212> DNA

<213> Artificial Sequence

<220>  
 <223> Mouse B9 heavy chain variable framework 3 (FW3) sequence  
  
 <400> 1552  
 cgattcacca tctccagaga caatgccaaag aacaccctgt acctgcaaat gagcagtctg 60  
 aagtctgagg acacagccat gtattactgt gca 93  
  
 <210> 1553  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B9 heavy chain variable framework 3 (FW3) sequence  
  
 <400> 1553  
  
 Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr Leu Gln  
 1 5 10 15  
  
 Met Ser Ser Leu Lys Ser Glu Asp Thr Ala Met Tyr Tyr Cys Ala Arg  
 20 25 30  
  
 <210> 1554  
 <211> 42  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B9 heavy chain variable complementarity determining regions  
 3 (CDR3) sequence  
  
 <400> 1554  
 agggataact acggtagtag ctacgactat gctatggact ac 42  
  
 <210> 1555  
 <211> 13  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B9 heavy chain variable complementarity determining regions  
 3 (CDR3) sequence  
  
 <400> 1555  
  
 Asp Asn Tyr Gly Ser Ser Tyr Asp Tyr Ala Met Asp Tyr  
 1 5 10  
  
 <210> 1556  
 <211> 33  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B9 heavy chain variable framework 4 (FW4) sequence  
  
 <400> 1556  
 tgggggtcaag gaacctcagt caccgtctcc tct 33  
  
 <210> 1557

<211> 11  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B9 heavy chain variable framework 4 (FW4) sequence

<400> 1557

Trp Gly Gln Gly Thr Ser Val Thr Val Ser Ser  
1 5 10

<210> 1558  
<211> 69  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B9 light chain variable framework 1 (FR1) sequence

<400> 1558  
caaattgttc tcaccagtc tccagcaatc atgtctgcat ctccagggga ggaggtcacc 60  
ctaacctgc 69

<210> 1559  
<211> 23  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B9 light chain variable framework 1 (FR1) sequence

<400> 1559

Gln Ile Val Leu Thr Gln Ser Pro Ala Ile Met Ser Ala Ser Pro Gly  
1 5 10 15

Glu Glu Val Thr Leu Thr Cys  
20

<210> 1560  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B9 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1560  
agtgccagct caagtgtaag ttacatgcac 30

<210> 1561  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B9 light chain variable complementarity determining regions  
1 (CDR1) sequence

<400> 1561



Ser Ala Ser Ser Ser Val Ser Tyr Met His  
1 5 10

<210> 1562  
<211> 45  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B9 light chain variable framework 2 (FR2) sequence

<400> 1562  
tgggtccagc agaggccagg cacttctccc aaactctgga tttat 45

<210> 1563  
<211> 15  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B9 light chain variable framework 2 (FR2) sequence

<400> 1563

Trp Phe Gln Gln Arg Pro Gly Thr Ser Pro Lys Leu Trp Ile Tyr  
1 5 10 15

<210> 1564  
<211> 21  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B9 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1564  
accacatcca acctggcttc t 21

<210> 1565  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse B9 light chain variable complementarity determining regions  
2 (CDR2) sequence

<400> 1565

Thr Thr Ser Asn Leu Ala Ser  
1 5

<210> 1566  
<211> 96  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Mouse B9 light chain variable framework 3 (FR3) sequence

<400> 1566  
ggagtccctg ctcgcttcag tggcagtgga tctgggacct cttactctct cacaatcagc 60

cgaatggagg ctgaagatgc tgccacttat tactgc

96

<210> 1567

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B9 light chain variable framework 3 (FR3) sequence

<400> 1567

Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Thr Ser Tyr Ser  
1 5 10 15

Leu Thr Ile Ser Arg Met Glu Ala Glu Asp Ala Ala Thr Tyr Tyr Cys  
20 25 30

<210> 1568

<211> 24

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B9 light chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 1568

cagcaaagga gtagttaccc attc

24

<210> 1569

<211> 8

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B9 light chain variable complementarity determining regions  
3 (CDR3) sequence

<400> 1569

Gln Gln Arg Ser Ser Tyr Pro Phe  
1 5

<210> 1570

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Mouse B9 light chain variable framework 4 (FR4) sequence

<400> 1570

acgttcggct cggggacaaa gttggaaata aaa

33

<210> 1571

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Mouse B9 light chain variable framework 4 (FR4) sequence

<400> 1571

Thr Phe Gly Ser Gly Thr Lys Leu Glu Ile Lys  
1 5 10

<210> 1572

<211> 741

<212> DNA

<213> Artificial Sequence

<220>

<223> mu3C2B1 scFv sequence

<400> 1572

gaagtgatgc tggaggagtc tgggggaggc ttagtgaagc ctggagggtc cctgaaactc 60  
tcctgtgcag cctctggaat cactttcagt acctatacca tgctgtgggt tcgccagact 120  
ccggagaaga ggctggagtg ggtcgcaacc attagtactg gtggtgataa aacctactat 180  
tcagacagtg tgaagggtcg attcaccatc tccagagaca atgccaagaa caacctgtac 240  
ctccaaatga gcagtctgag gtctgaggac acggccttgt attactgtgc aaggggaacc 300  
acggctatgt attactatgc tatggactac tggggtaag gaacctcagt caccgtctcc 360  
tcagggtggcg gaggatctgg cggaggtgga agcggcggag gcggatccga cattgtgctg 420  
acacagtctc ctgcttcctt agctgtatct ctggggcaga gggccaccat ctcatgcagg 480  
gccagcaaaa gtatcagtac atctgactat aattatattc actggtacca acagaaacca 540  
ggacagccac ccaaactcct catctatctt gcatccaacc tagaatctgg ggtccctgcc 600  
aggttcagtg gcagtgggtc tgggacagac ttcacctca acatccatcc tgtggaggaa 660  
gaagatgctg caacctatta ctgtcagcac agtagggagc ttcctctcac gttcgggtgct 720  
gggaccaagc tggagctgaa a 741

<210> 1573

<211> 247

<212> PRT

<213> Artificial Sequence

<220>

<223> mu3C2B1 scFv sequence

<400> 1573

Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Ile Thr Phe Ser Thr Tyr  
20 25 30

Thr Met Ser Trp Val Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val  
35 40 45

Ala Thr Ile Ser Thr Gly Gly Asp Lys Thr Tyr Tyr Ser Asp Ser Val  
50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Asn Leu Tyr

65	70	75	80
Leu Gln Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Leu Tyr Tyr Cys	85	90	95
Ala Arg Gly Thr Thr Ala Met Tyr Tyr Tyr Ala Met Asp Tyr Trp Gly	100	105	110
Gln Gly Thr Ser Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly	115	120	125
Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Leu Thr Gln Ser Pro	130	135	140
Ala Ser Leu Ala Val Ser Leu Gly Gln Arg Ala Thr Ile Ser Cys Arg	145	150	155
Ala Ser Lys Ser Ile Ser Thr Ser Asp Tyr Asn Tyr Ile His Trp Tyr	165	170	175
Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Leu Ala Ser	180	185	190
Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly	195	200	205
Thr Asp Phe Thr Leu Asn Ile His Pro Val Glu Glu Glu Asp Ala Ala	210	215	220
Thr Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro Leu Thr Phe Gly Ala	225	230	235
			240
Gly Thr Lys Leu Glu Leu Lys	245		

<210> 1574

<211> 741

<212> DNA

<213> Artificial Sequence

<220>

<223> mu20A10 scFv full sequence

<400> 1574

gaagtgatgc tgggtggaatc tggcggcgga ctggttaagc ctggcggatc tctgaagctg	60
agctgtgccg ccagcggcctt caccttttagc acatacgcca tgagctggat ccggcagacc	120
cctgagaaga gactggaatg ggttgccagc atcggcagag ccggcagcac ctactacagc	180
gattctgtga agggcagatt caccatcagc cgggacaacg tgcggaacat cctgtacctg	240
cagatgagca gcctgcggag cgaggatacc gccatgtact actgtgccag aggacccatc	300
tacaacgact acgacgagtt cgcctattgg ggccagggca cactggttac agtttctgct	360
ggtggcggag gatctggcgg aggtggaagc ggcggaggcg gatccaatat catgatgaca	420

cagagcccca gcagcctggc tgtgtctgct ggcgagaaag tgaccatgtc ctgcaagagc	480
agccagagcg tgctgtactc cagcaaccag aagaactacc tggcctggta tcagcagaag	540
cccgccagct ctccaaagct gctgatctac tgggccagca ccagagaaag cggcgtgccc	600
gatagattca caggcagcgg cagcggaacc gacttcaccc tgacaatcag ctctgtgcag	660
gccgaagatc tggccgtgta ctattgccac cagtacctgt ccagcctgac ctttggcgcc	720
ggaacaaagc tggaactgaa g	741

<210> 1575  
 <211> 247  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> mu20A10 scFv full sequence  
  
 <400> 1575

Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly	1	5	10	15
Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Thr Tyr	20	25	30	
Ala Met Ser Trp Ile Arg Gln Thr Pro Glu Lys Arg Leu Glu Trp Val	35	40	45	
Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr Ser Asp Ser Val Lys	50	55	60	
Gly Arg Phe Thr Ile Ser Arg Asp Asn Val Arg Asn Ile Leu Tyr Leu	65	70	75	80
Gln Met Ser Ser Leu Arg Ser Glu Asp Thr Ala Met Tyr Tyr Cys Ala	85	90	95	
Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe Ala Tyr Trp Gly Gln	100	105	110	
Gly Thr Leu Val Thr Val Ser Ala Gly Gly Gly Gly Ser Gly Gly Gly	115	120	125	
Gly Ser Gly Gly Gly Gly Ser Asn Ile Met Met Thr Gln Ser Pro Ser	130	135	140	
Ser Leu Ala Val Ser Ala Gly Glu Lys Val Thr Met Ser Cys Lys Ser	145	150	155	160
Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys Asn Tyr Leu Ala Trp	165	170	175	
Tyr Gln Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr Trp Ala				

180

185

190

Ser Thr Arg Glu Ser Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser  
195 200 205

Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Val Gln Ala Glu Asp Leu  
210 215 220

Ala Val Tyr Tyr Cys His Gln Tyr Leu Ser Ser Leu Thr Phe Gly Ala  
225 230 235 240

Gly Thr Lys Leu Glu Leu Lys  
245

&lt;210&gt; 1576

&lt;211&gt; 741

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; hu20A10M scFV

&lt;400&gt; 1576

gaggtgcagc tggttgaatc tggcggcgga cttgtgaagc ctggcggatc tctgagactg 60

agctgtgccg ccagcggcctt caccttttagc acatacgcca tgagctgggt ccgacaggcc 120

cctggaaaag gccttgaatg ggttgccctc atcggcagag ccggcagcac ctactacagc 180

gattctgtga agggcagatt caccatcagc cgggacaacg ccaagaacag cctgtacctg 240

cagatgaact ccctgagagc cgaggacacc gccgtgtact attgtgccag aggacccatc 300

tacaacgact acgacgagtt gcctatttgg ggccagggca cactgggtcac agtcagctct 360

ggcgggtggcg gaagcggagg cgggtggctcc ggtggcggag gcagcgacat cgtgatgaca 420

cagagccctt ctagcctggc cgtgtctctg ggagagagag ccacaatcag ctgcaagagc 480

agccagagcg tgctgtactc cagcaaccag aagaactacc tggcctggta tcagcagaag 540

cccggacagt ctcccaagct gctgatctac tgggccagca ccagagaaaag cggcgtgccc 600

gatagattca caggctctgg cagcggcacc gacttcaccc tgacaattag cagtctgcag 660

gccgaggacg tggccgtgta ctactgtcac cagtacctga gcagcctgac ctttggcggc 720

ggaacaaagg tggaaatcaa g 741

&lt;210&gt; 1577

&lt;211&gt; 247

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; hu20A10M scFV

&lt;400&gt; 1577

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Thr Tyr  
 20 25 30  
 Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
 35 40 45  
 Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr Ser Asp Ser Val Lys  
 50 55 60  
 Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr Leu  
 65 70 75 80  
 Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys Ala  
 85 90 95  
 Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe Ala Tyr Trp Gly Gln  
 100 105 110  
 Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly  
 115 120 125  
 Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser Pro Ser  
 130 135 140  
 Ser Leu Ala Val Ser Leu Gly Glu Arg Ala Thr Ile Ser Cys Lys Ser  
 145 150 155 160  
 Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys Asn Tyr Leu Ala Trp  
 165 170 175  
 Tyr Gln Gln Lys Pro Gly Gln Ser Pro Lys Leu Leu Ile Tyr Trp Ala  
 180 185 190  
 Ser Thr Arg Glu Ser Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser  
 195 200 205  
 Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Leu Gln Ala Glu Asp Val  
 210 215 220  
 Ala Val Tyr Tyr Cys His Gln Tyr Leu Ser Ser Leu Thr Phe Gly Gly  
 225 230 235 240  
 Gly Thr Lys Val Glu Ile Lys  
 245

<210> 1578

<211> 744

<212> DNA

<213> Artificial Sequence

<220>

<223> hu20A10C2 scFV

<400> 1578

gaggtgcagc tggttgaatc tggcggcgga cttgtgaagc ctggcggatc tctgagactg 60  
agctgtgccg ccagcggctt caccttttagc acatacgcca tgagctgggt ccgacaggcc 120  
cctggaaaag gccttgaatg ggttgacctt atcggcagag ccggcagcac ctactacagc 180  
gattctgtga agggcagatt caccatcagc cgggacaacg ccaagaacag cctgtacctg 240  
cagatgaact ccctgagagc cgaggacacc gccgtgtact attgtgccag aggacccatc 300  
tacaacgact acgacgagtt cgcctattgg ggccagggca cactgggtcac agtcagctct 360  
ggcgggtggcg gaagcggagg cgggtggctcc ggtggcggag gcagcgacat tgtgctgacc 420  
cagtctccag cctccttggc cgtgtctcca ggacagaggg ccaccatcac ctgcaagagc 480  
agccagagcg tgctgtactc cagcaaccag aagaactacc tggcctggta tcagcagaaa 540  
ccaggacaac ctctaaact cctgatttac tgggccagca ccagagaaaag cgggggtcca 600  
gccaggttca gcggcagtggt gtctgggacc gatttcaccc tcacaattaa tcctgtggaa 660  
gctaatagata ctgcaattaa ttactgtcac cagtacctga gcagcctgac cttcggcgga 720  
gggaccaagg tggagatcaa acga 744

<210> 1579  
<211> 248  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> hu20A10C2 scFV

<400> 1579

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15  
  
Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Thr Tyr  
20 25 30  
  
Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45  
  
Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr Ser Asp Ser Val Lys  
50 55 60  
  
Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr Leu  
65 70 75 80  
  
Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys Ala  
85 90 95  
  
Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe Ala Tyr Trp Gly Gln  
100 105 110  
  
Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly  
115 120 125



Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Leu Thr Gln Ser Pro Ala  
130 135 140

Ser Leu Ala Val Ser Pro Gly Gln Arg Ala Thr Ile Thr Cys Lys Ser  
145 150 155 160

Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys Asn Tyr Leu Ala Trp  
165 170 175

Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Trp Ala  
180 185 190

Ser Thr Arg Glu Ser Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser  
195 200 205

Gly Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu Ala Asn Asp Thr  
210 215 220

Ala Asn Tyr Tyr Cys His Gln Tyr Leu Ser Ser Leu Thr Phe Gly Gly  
225 230 235 240

Gly Thr Lys Val Glu Ile Lys Arg  
245

<210> 1580  
<211> 744  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> hu20A10N scFV

<400> 1580  
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cctggcaaag gcctggaatg ggtggcgtct attggcagag ccggcagcac ctactacagc 180  
gactctgtga agggcagatt caccatcagc cgggacaacg ccaagaacag cctgtacctg 240  
cagatgaact ccctgagagc cgaggacacc gccgtgtact attgtgccag aggacccatc 300  
tacaacgact acgacgagtt cgcctattgg ggccagggca cactgggtcac agtttctagc 360  
ggcgggtggcg gaagcggagg cgggtggctcc ggtggcggag gcagcgaaat tgtgctgaca 420  
cagagccccg ccacactgtc actttctcca ggcgaaagag ccacactgag ctgcaagagc 480  
agccagagcg tgctgtactc cagcaaccag aagaactacc tggcctggta tcagcagaag 540  
cccggccaag ctctctggct gctgatctat tgggccagca caagagagag cggcatccct 600  
gccagatttt ctggcagcgg ctctggcacc gatttcaccc tgaccataag cagcctggaa 660  
cctgaggact tcgccgtgta ttactgccac cagtacctga gcagcctgac ctttggcgga 720  
ggcaccaagg tggaaatcaa gcgg 744

<210> 1581

<211> 248  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> hu20A10N scFV

<400> 1581

Gln Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Thr Tyr  
20 25 30

Ala Met Ser Trp Ile Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45

Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr Ser Asp Ser Val Lys  
50 55 60

Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr Leu  
65 70 75 80

Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys Ala  
85 90 95

Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe Ala Tyr Trp Gly Gln  
100 105 110

Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly  
115 120 125

Gly Ser Gly Gly Gly Gly Ser Glu Ile Val Leu Thr Gln Ser Pro Ala  
130 135 140

Thr Leu Ser Leu Ser Pro Gly Glu Arg Ala Thr Leu Ser Cys Lys Ser  
145 150 155 160

Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys Asn Tyr Leu Ala Trp  
165 170 175

Tyr Gln Gln Lys Pro Gly Gln Ala Pro Arg Leu Leu Ile Tyr Trp Ala  
180 185 190

Ser Thr Arg Glu Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser  
195 200 205

Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe  
210 215 220

Ala Val Tyr Tyr Cys His Gln Tyr Leu Ser Ser Leu Thr Phe Gly Gly  
225 230 235 240

Gly Thr Lys Val Glu Ile Lys Arg  
245

<210> 1582  
<211> 1479  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> mu20A10CAR T841BB3z

<400> 1582  
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ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg gatccggcag 180  
acccctgaga agagactgga atgggttgcc agcatcggca gagccggcag cacctactac 240  
agcgattctg tgaagggcag attcaccatc agccgggaca acgtgcggaa catcctgtac 300  
ctgcagatga gcagcctgcg gagcgaggat accgccatgt actactgtgc cagaggaccc 360  
atctacaacg actacgacga gttcgcctat tggggccagg gcacactggt tacagtttct 420  
gctggtggcg gaggatctgg cggaggtgga agcggcggag gcggatcaa tatcatgatg 480  
acacagagcc cgagcagcct ggctgtgtct gctggcgaga aagtgaccat gtcctgcaag 540  
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag 600  
aagcccggcc agtctcctaa gctgctgata tactgggccg gcaccagaga aagcggcgtg 660  
cccgatagat tcacaggcag cggcagcggc accgacttca ccctgacaat cagctctgtg 720  
caggccgaag atctggccgt gtactattgc caccagtacc tgtccagcct gacctttggc 780  
gccggaacaa agctggaact gaagacaaca acccctgccc ccagacctcc taccagcc 840  
cctacaattg ccagccagcc tctgagcctg agggccgagg cttgtagacc tgctgctggc 900  
ggagccgtgc acaccagagg actggatttc gcctgcgaca tctacatctg ggcgcccttg 960  
gccgggactt gtggggctct tctcctgtca ctggttatca ccctttactg caaacggggc 1020  
agaaagaaac tcctgtatat attcaaaaaa ccatttatga gaccagtaca aactactcaa 1080  
gaggaagatg gctgtagctg ccgatttcca gaagaagaag aaggaggatg tgaactgaga 1140  
gtgaagttca gcaggagcgc agacgcccc gcgtacaagc agggccagaa ccagctctat 1200  
aacgagctca atctaggacg aagagaggag tacgatgttt tggacaagag acgtggccgg 1260  
gacctgaga tggggggaaa gccgagaagg aagaaccctc aggaaggcct gtacaatgaa 1320  
ctgcagaaag ataagatggc ggaggcctac agtgagattg ggatgaaagg cgagcgccgg 1380  
aggggcaagg ggacgatgg cttttaccag ggtctcagta cagccaccaa ggacacctac 1440  
gacgcccttc acatgcaggc cctgccccct cgctgataa 1479

<210> 1583  
<211> 491  
<212> PRT  
<213> Artificial Sequence

<220>

<223> mu20A10CAR T841BB3z

<400> 1583

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Thr Tyr Ala Met Ser Trp Ile Arg Gln Thr Pro Glu Lys  
50 55 60

Arg Leu Glu Trp Val Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr  
65 70 75 80

Ser Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Val Arg  
85 90 95

Asn Ile Leu Tyr Leu Gln Met Ser Ser Leu Arg Ser Glu Asp Thr Ala  
100 105 110

Met Tyr Tyr Cys Ala Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe  
115 120 125

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala Gly Gly Gly  
130 135 140

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asn Ile Met Met  
145 150 155 160

Thr Gln Ser Pro Ser Ser Leu Ala Val Ser Ala Gly Glu Lys Val Thr  
165 170 175

Met Ser Cys Lys Ser Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys  
180 185 190

Asn Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ser Pro Lys Leu  
195 200 205

Leu Ile Tyr Trp Ala Ser Thr Arg Glu Ser Gly Val Pro Asp Arg Phe  
210 215 220

Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Val  
225 230 235 240

Gln Ala Glu Asp Leu Ala Val Tyr Tyr Cys His Gln Tyr Leu Ser Ser  
245 250 255

Leu Thr Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys Thr Thr Thr Pro  
260 265 270

Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu  
275 280 285

Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His  
290 295 300

Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu  
305 310 315 320

Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr  
325 330 335

Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe  
340 345 350

Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg  
355 360 365

Phe Pro Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser  
370 375 380

Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr  
385 390 395 400

Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys  
405 410 415

Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn  
420 425 430

Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu  
435 440 445

Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly  
450 455 460

His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr  
465 470 475 480

Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1584  
<211> 1482  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> hu20A10CAR T841BB3z

<400> 1584

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ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg ggtccgacag 180  
gccccctggaa aaggccttga atgggttgcc tctatcggca gagccggcag cacctactac 240  
agcgattctg tgaagggcag attcaccatc agccgggaca acgccaagaa cagcctgtac 300  
ctgcagatga actccctgag agccgaggac accgccgtgt actattgtgc cagaggaccc 360  
atctacaacg actacgacga gttcgcctat tggggccagg gcacactggt cacagtcagc 420  
tctggcgggtg gcggaagcgg aggcgggtggc tccggtggcg gaggcagcga catttgtctg 480  
accagtcctc cagcctcctt ggccgtgtct ccaggacaga gggccaccat cacctgcaag 540  
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag 600  
aaaccaggac aacctcctaa actcctgatt tactgggccg gcaccagaga aagcggggtc 660  
ccagccagggt tcagcggcag tgggtctggg accgatttca ccctcacaat taatcctgtg 720  
gaagctaattg atactgcaaa ttattactgt caccagtacc tgagcagcct gaccttcggc 780  
ggaggggacca aggtggagat caaacgaaca acaaccctg cccccagacc tcctaccca 840  
gccccataaa ttgccagcca gcctctgagc ctgaggcccg aggcttgtag acctgtgct 900  
ggcggagccg tgcacaccag aggactggat ttgcctgcg acatctacat ctgggcgccc 960  
ttggccggga cttgtggggt ctttctcctg tctactggtta tcacccttta ctgcaaacgg 1020  
ggcagaaaga aactcctgta tatattcaaa caaccattta tgagaccagt acaaactact 1080  
caagaggaag atggctgtag ctgccgattt ccagaagaag aagaaggagg atgtgaactg 1140  
agagtgaagt tcagcaggag cgcagacgcc cccgcgtaca agcagggccg gaaccagctc 1200  
tataacgagc tcaatctagg acgaagagag gagtacgatg ttttggacaa gagacgtggc 1260  
cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgtacaat 1320  
gaactgcaga aagataagat ggcggaggcc tacagtgaga ttgggatgaa aggcgagcgc 1380  
cggaggggca aggggcacga tggcctttac cagggtctca gtacagccac caaggacacc 1440  
tacgacgcc ttcacatgca ggccctgccc cctcgctgat aa 1482

<210> 1585  
<211> 492  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> hu20A10CAR T841BB3z

<400> 1585

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Thr Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Gly Leu Glu Trp Val Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr  
65 70 75 80

Ser Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys  
85 90 95

Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala  
100 105 110

Val Tyr Tyr Cys Ala Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe  
115 120 125

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly  
130 135 140

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Leu  
145 150 155 160

Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala Thr  
165 170 175

Ile Thr Cys Lys Ser Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys  
180 185 190

Asn Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu  
195 200 205

Leu Ile Tyr Trp Ala Ser Thr Arg Glu Ser Gly Val Pro Ala Arg Phe  
210 215 220

Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Pro Val  
225 230 235 240

Glu Ala Asn Asp Thr Ala Asn Tyr Tyr Cys His Gln Tyr Leu Ser Ser  
245 250 255

Leu Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr  
260 265 270

Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro  
275 280 285

Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val  
290 295 300

His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro

305		310		315		320
Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu						
		325		330		335
Tyr Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro						
		340		345		350
Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys						
		355		360		365
Arg Phe Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe						
		370		375		380
Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu						
		385		390		395
Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp						
		405		410		415
Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys						
		420		425		430
Asn Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala						
		435		440		445
Glu Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys						
		450		455		460
Gly His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr						
		465		470		475
				480		
Tyr Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg						
		485		490		

<210> 1586  
 <211> 1476  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> mu20A10CAR T8283z

<400> 1586	
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ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg gatccggcag	180
acccttgaga agagactgga atgggttgcc agcatcggca gagccggcag cacctactac	240
agcgattctg tgaagggcag attcaccatc agccgggaca acgtgcggaa catcctgtac	300
ctgcagatga gcagcctgcg gagcgaggat accgccatgt actactgtgc cagaggaccc	360
atctacaacg actacgacga gttcgcctat tggggccagg gcacactggt tacagtttct	420



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acacagagcc cgagcagcct ggctgtgtct gctggcgaga aagtgaccat gtcctgcaag 540  
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag 600  
aagcccggcc agtctcctaa gctgctgata tactggggcca gcaccagaga aagcggcgtg 660  
cccgatagat tcacaggcag cggcagcgga accgacttca ccctgacaat cagctctgtg 720  
caggccgaag atctggccgt gtactattgc caccagtacc tgtccagcct gacctttggc 780  
gccggaacaa agctggaact gaagacaaca acccctgccc ccagacctcc taccagcc 840  
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gccgggactt gtggggctct tctcctgtca ctggttatca ccctttactg caggagtaag 1020  
aggagcaggc tcctgcacag tgactacatg aacatgactc ctagaagacc tgggcctacc 1080  
agaaagcatt accagcccta tgccccacca cgcgacttcg cagcctatcg ctccagagtg 1140  
aagttcagca ggagcgcaga cggcccgcg tacaagcagg gccagaacca gctctataac 1200  
gagctcaatc taggacgaag agaggagtac gatgttttgg acaagagacg tggccgggac 1260  
cctgagatgg ggggaaagcc gagaaggaag aaccctcagg aaggcctgta caatgaactg 1320  
cagaaagata agatggcgga ggccctacagt gagattggga tgaaaggcga gcgccggagg 1380  
ggcaaggggc acgatggcct ttaccagggt ctacgtacag ccaccaagga cacctacgac 1440  
gcccttcaca tgcaggccct gcccctcgc tgataa 1476

<210> 1587  
<211> 490  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> mu20A10CAR T8283z

<400> 1587

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15  
  
His Ala Ala Arg Pro Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30  
  
Val Lys Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45  
  
Thr Phe Ser Thr Tyr Ala Met Ser Trp Ile Arg Gln Thr Pro Glu Lys  
50 55 60  
  
Arg Leu Glu Trp Val Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr  
65 70 75 80  
  
Ser Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Val Arg

85

90

95

Asn Ile Leu Tyr Leu Gln Met Ser Ser Leu Arg Ser Glu Asp Thr Ala  
 100 105 110

Met Tyr Tyr Cys Ala Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe  
 115 120 125

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala Gly Gly Gly  
 130 135 140

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Ser Asn Ile Met Met  
 145 150 155 160

Thr Gln Ser Pro Ser Ser Leu Ala Val Ser Ala Gly Glu Lys Val Thr  
 165 170 175

Met Ser Cys Lys Ser Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys  
 180 185 190

Asn Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ser Pro Lys Leu  
 195 200 205

Leu Ile Tyr Trp Ala Ser Thr Arg Glu Ser Gly Val Pro Asp Arg Phe  
 210 215 220

Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Val  
 225 230 235 240

Gln Ala Glu Asp Leu Ala Val Tyr Tyr Cys His Gln Tyr Leu Ser Ser  
 245 250 255

Leu Thr Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys Thr Thr Thr Pro  
 260 265 270

Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu  
 275 280 285

Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His  
 290 295 300

Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu  
 305 310 315 320

Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr  
 325 330 335

Cys Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met  
 340 345 350

Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala  
 355 360 365

Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser Arg  
370 375 380

Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn  
385 390 395 400

Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg  
405 410 415

Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro  
420 425 430

Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala  
435 440 445

Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His  
450 455 460

Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr Asp  
465 470 475 480

Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1588  
<211> 1479  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> hu20A10CAR T8283z

<400> 1588  
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ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg ggtccgacag 180  
gccccctggaa aaggccttga atgggttgcc tctatcggca gagccggcag cacctactac 240  
agcgattctg tgaagggcag attcaccatc agccgggaca acgccaagaa cagcctgtac 300  
ctgcagatga actccctgag agccgaggac accgccgtgt actattgtgc cagaggaccc 360  
atctacaacg actacgacga gttcgcctat tggggccagg gcacactggt cacagtcagc 420  
tctggcgggtg gcggaagcgg aggcggtggc tccggtggcg gaggcagcga cattgtgctg 480  
accagttctc cagcctcctt ggccgtgtct ccaggacaga gggccacat cacctgcaag 540  
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag 600  
aaaccaggac aacctcctaa actcctgatt tactgggccg gcaccagaga aagcggggtc 660  
ccagccaggt tcagcggcag tgggtctggg accgatttca ccctcacaat taatcctgtg 720  
gaagctaatt atactgcaaa ttattactgt caccagtacc tgagcagcct gaccttcggc 780

ggagggacca aggtggagat caaacgaaca acaaccctg ccccgagacc tcctaccca 840  
gcccctacaa ttgccagcca gcctctgagc ctgaggcccg aggcttgtag acctgctgct 900  
ggcggagccg tgcacaccag aggactggat ttgcctgcg acatctacat ctgggcgccc 960  
ttggccggga cttgtggggt ctttctcctg tcaactgtta tcacccttta ctgcaggagt 1020  
aagaggagca ggctcctgca cagtgactac atgaacatga ctcctagaag acctgggcct 1080  
accagaaagc attaccagcc ctatgcccc ccacgcgact tcgcagccta tcgctccaga 1140  
gtgaagtcca gcaggagcgc agacgcccc gcgtacaagc agggccagaa ccagctctat 1200  
aacgagctca atctaggacg aagagaggag tacgatgttt tggacaagag acgtggccgg 1260  
gaccctgaga tggggggaaa gccgagaagg aagaaccctc aggaaggcct gtacaatgaa 1320  
ctgcagaaag ataagatggc ggaggcctac agtgagattg ggatgaaagg cgagcgccgg 1380  
aggggcaagg ggacagatgg cttttaccag ggtctcagta cagccaccaa ggacacctac 1440  
gacgcccttc acatgcaggc cctgccccct cgctgataa 1479

<210> 1589  
<211> 491  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> hu20A10CAR T8283z

<400> 1589

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15  
  
His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30  
  
Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45  
  
Thr Phe Ser Thr Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60  
  
Gly Leu Glu Trp Val Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr  
65 70 75 80  
  
Ser Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys  
85 90 95  
  
Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala  
100 105 110  
  
Val Tyr Tyr Cys Ala Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe  
115 120 125  
  
Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly  
130 135 140

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Leu  
145 150 155 160

Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala Thr  
165 170 175

Ile Thr Cys Lys Ser Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys  
180 185 190

Asn Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu  
195 200 205

Leu Ile Tyr Trp Ala Ser Thr Arg Glu Ser Gly Val Pro Ala Arg Phe  
210 215 220

Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Pro Val  
225 230 235 240

Glu Ala Asn Asp Thr Ala Asn Tyr Tyr Cys His Gln Tyr Leu Ser Ser  
245 250 255

Leu Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr  
260 265 270

Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro  
275 280 285

Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val  
290 295 300

His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro  
305 310 315 320

Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu  
325 330 335

Tyr Cys Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn  
340 345 350

Met Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr  
355 360 365

Ala Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser  
370 375 380

Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr  
385 390 395 400

Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys  
405 410 415

Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn  
420 425 430

Pro Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu  
435 440 445

Ala Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly  
450 455 460

His Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr  
465 470 475 480

Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1590

<211> 1479

<212> DNA

<213> Artificial Sequence

<220>

<223> mu20A10CAR T841BB3z1XX

<400> 1590

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ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg gatccggcag	180
acccctgaga agagactgga atgggttgcc agcatcggca gagccggcag cacctactac	240
agcgattctg tgaagggcag attcaccatc agccgggaca acgtgcggaa catcctgtac	300
ctgcagatga gcagcctgcg gagcgaggat accgccatgt actactgtgc cagaggaccc	360
atctacaacg actacgacga gttcgcctat tggggccagg gcacactggt tacagtttct	420
gctggtggcg gaggatctgg cggaggtgga agcggcggag gcggatcaa tatcatgatg	480
acacagagcc cgagcagcct ggctgtgtct gctggcgaga aagtgaccat gtcctgcaag	540
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag	600
aagcccggcc agtctcctaa gctgctgata tactgggccca gcaccagaga aagcggcgtg	660
cccgatagat tcacaggcag cggcagcggc accgacttca ccctgacaat cagctctgtg	720
caggccgaag atctggccgt gtactattgc caccagtacc tgtccagcct gacctttggc	780
gccggaacaa agctggaact gaagacaaca acccctgccc ccagacctcc taccagcc	840
cctacaattg ccagccagcc tctgagcctg agggccgagg cttgtagacc tgctgctggc	900
ggagccgtgc acaccagagg actggatttc gcctgcgaca tctacatctg ggcgcccttg	960
gccgggactt gtggggctct tctcctgtca ctggttatca ccctttactg caaacggggc	1020
agaaagaac tcctgtatat attcaaaaa ccatttatga gaccagtaca aactactcaa	1080
gaggaagatg gctgtagctg ccgatttcca gaagaagaag aaggaggatg tgaactgaga	1140
gtgaagttca gcaggagcgc agacgcccc gcgtacaagc agggccagaa ccagctctat	1200

aacgagctca atctaggacg aagagaggag tacgatgttt tggacaagag acgtggccgg	1260
gaccttgaga tgggggggaaa gccgagaagg aagaaccctc aggaaggcct gttcaatgaa	1320
ctgcagaaag ataagatggc ggaggccttc agtgagattg ggatgaaagg cgagcgccgg	1380
aggggcaagg ggcacgatgg ccttttccag ggtctcagta cagccaccaa ggacaccttc	1440
gacgcccttc acatgcaggc cctgccccct cgctgataa	1479

<210> 1591  
 <211> 491  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> mu20A10CAR T841BB3z1XX  
  
 <400> 1591

Met	Ala	Leu	Pro	Val	Thr	Ala	Leu	Leu	Leu	Pro	Leu	Ala	Leu	Leu	Leu	1	5	10	15
His	Ala	Ala	Arg	Pro	Glu	Val	Met	Leu	Val	Glu	Ser	Gly	Gly	Gly	Leu	20	25	30	
Val	Lys	Pro	Gly	Gly	Ser	Leu	Lys	Leu	Ser	Cys	Ala	Ala	Ser	Gly	Phe	35	40	45	
Thr	Phe	Ser	Thr	Tyr	Ala	Met	Ser	Trp	Ile	Arg	Gln	Thr	Pro	Glu	Lys	50	55	60	
Arg	Leu	Glu	Trp	Val	Ala	Ser	Ile	Gly	Arg	Ala	Gly	Ser	Thr	Tyr	Tyr	65	70	75	80
Ser	Asp	Ser	Val	Lys	Gly	Arg	Phe	Thr	Ile	Ser	Arg	Asp	Asn	Val	Arg	85	90	95	
Asn	Ile	Leu	Tyr	Leu	Gln	Met	Ser	Ser	Leu	Arg	Ser	Glu	Asp	Thr	Ala	100	105	110	
Met	Tyr	Tyr	Cys	Ala	Arg	Gly	Pro	Ile	Tyr	Asn	Asp	Tyr	Asp	Glu	Phe	115	120	125	
Ala	Tyr	Trp	Gly	Gln	Gly	Thr	Leu	Val	Thr	Val	Ser	Ala	Gly	Gly	Gly	130	135	140	
Gly	Ser	Gly	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Ser	Asn	Ile	Met	Met		145	150	155	160
Thr	Gln	Ser	Pro	Ser	Ser	Leu	Ala	Val	Ser	Ala	Gly	Glu	Lys	Val	Thr	165	170	175	
Met	Ser	Cys	Lys	Ser	Ser	Gln	Ser	Val	Leu	Tyr	Ser	Ser	Asn	Gln	Lys	180	185	190	

Asn Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ser Pro Lys Leu  
 195 200 205

Leu Ile Tyr Trp Ala Ser Thr Arg Glu Ser Gly Val Pro Asp Arg Phe  
 210 215 220

Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Val  
 225 230 235 240

Gln Ala Glu Asp Leu Ala Val Tyr Tyr Cys His Gln Tyr Leu Ser Ser  
 245 250 255

Leu Thr Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys Thr Thr Thr Pro  
 260 265 270

Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu  
 275 280 285

Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His  
 290 295 300

Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu  
 305 310 315 320

Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr  
 325 330 335

Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe  
 340 345 350

Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg  
 355 360 365

Phe Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser  
 370 375 380

Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr  
 385 390 395 400

Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys  
 405 410 415

Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn  
 420 425 430

Pro Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu  
 435 440 445

Ala Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly  
 450 455 460



His Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe  
 465 470 475 480

Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
 485 490

<210> 1592  
 <211> 1482  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> hu20A10CAR T841BB3z1XX

<400> 1592  
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 ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg ggtccgacag 180  
 gcccctggaa aaggccttga atgggttgcc tctatcggca gagccggcag cacctactac 240  
 agcgattctg tgaagggcag attcaccatc agccgggaca acgccaagaa cagcctgtac 300  
 ctgcagatga actccctgag agccgaggac accgccgtgt actattgtgc cagaggaccc 360  
 atctacaacg actacgacga gttcgcttat tggggccagg gcacactggt cacagtcagc 420  
 tctggcgggtg gcggaagcgg aggcgggtggc tccggtggcg gaggcagcga catttgtctg 480  
 acccagtctc cagcctcctt ggccgtgtct ccaggacaga gggccaccat cacctgcaag 540  
 agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag 600  
 aaaccaggac aacctcctaa actcctgatt tactgggcca gcaccagaga aagcggggtc 660  
 ccagccaggt tcagcggcag tgggtctggg accgatttca ccctcacaat taatcctgtg 720  
 gaagctaatt atactgcaaa ttattactgt caccagtacc tgagcagcct gaccttcggc 780  
 ggaggggacca aggtggagat caaacgaaca acaaccctg ccccagacc tcctaccca 840  
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 ggccggagccg tgcacaccag aggactggat ttgcctgag acatctacat ctgggcgccc 960  
 ttggccggga cttgtgggtt ctttctcctg tctactggtt tcacccttta ctgcaaacgg 1020  
 ggagaaaga aactcctgta tatattcaaa caaccattta tgagaccagt acaaactact 1080  
 caagaggaag atggctgtag ctgccgattt ccagaagaag aagaaggagg atgtgaactg 1140  
 agagtgaagt tcagcaggag cgcagacgcc cccgcgtaca agcagggccga gaaccagctc 1200  
 tataacgagc tcaatctagg acgaagagag gagtacgatg ttttgacaa gagacgtggc 1260  
 cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgttcaat 1320  
 gaactgcaga aagataagat ggccggaggcc ttcagtgaga ttgggatgaa aggcgagcgc 1380  
 cggaggggca aggggcacga tggccttttc cagggtctca gtacagccac caaggacacc 1440  
 ttcgacgcc ttcacatgca ggccctgccc cctcgctgat aa 1482

<210> 1593

<211> 492  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> hu20A10CAR T841BB3z1XX

<400> 1593

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Thr Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Gly Leu Glu Trp Val Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr  
65 70 75 80

Ser Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys  
85 90 95

Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala  
100 105 110

Val Tyr Tyr Cys Ala Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe  
115 120 125

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly  
130 135 140

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Leu  
145 150 155 160

Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala Thr  
165 170 175

Ile Thr Cys Lys Ser Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys  
180 185 190

Asn Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu  
195 200 205

Leu Ile Tyr Trp Ala Ser Thr Arg Glu Ser Gly Val Pro Ala Arg Phe  
210 215 220

Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Pro Val  
225 230 235 240

Glu Ala Asn Asp Thr Ala Asn Tyr Tyr Cys His Gln Tyr Leu Ser Ser  
245 250 255

Leu Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr  
260 265 270

Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro  
275 280 285

Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val  
290 295 300

His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro  
305 310 315 320

Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu  
325 330 335

Tyr Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro  
340 345 350

Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys  
355 360 365

Arg Phe Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe  
370 375 380

Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu  
385 390 395 400

Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp  
405 410 415

Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys  
420 425 430

Asn Pro Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala  
435 440 445

Glu Ala Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys  
450 455 460

Gly His Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr  
465 470 475 480

Phe Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1594

<211> 1476

<212> DNA

<213> Artificial Sequence

<220>

<223> mu20A10CAR T8283z1XX

<400> 1594

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ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg gatccggcag	180
acccttgaga agagactgga atgggttgcc agcatcggca gagccggcag cacctactac	240
agcgattctg tgaagggcag attcaccatc agccgggaca acgtgcggaa catcctgtac	300
ctgcagatga gcagcctgcg gagcgaggat accgccatgt actactgtgc cagaggaccc	360
atctacaacg actacgacga gtctgcctat tggggccagg gcacactggt tacagtttct	420
gctggtggcg gaggatctgg cggaggtgga agcggcggag gcggatcaa tatcatgatg	480
acacagagcc cgagcagcct ggctgtgtct gctggcgaga aagtgacat gtcctgcaag	540
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cccgatagat tcacaggcag cggcagcggg accgacttca ccctgacaat cagctctgtg	720
caggccgaag atctggccgt gtactattgc caccagtacc tgtccagcct gacctttggc	780
gccggaacaa agctggaact gaagacaaca acccctgccc ccagacctcc taccagcc	840
cctacaattg ccagccagcc tctgagcctg agggccgagg cttgtagacc tgctgctggc	900
ggagccgtgc acaccagagg actggatttc gcctgcgaca tctacatctg ggcgcccttg	960
gccgggactt gtggggctct tctcctgtca ctggttatca ccctttactg caggagtaag	1020
aggagcaggc tcctgcacag tgactacatg aacatgactc ctagaagacc tgggcctacc	1080
agaaagcatt accagcccta tgccccacca cgcgacttcg cagcctatcg ctccagagtg	1140
aagttcagca ggagcgcaga cggccccgcg tacaagcagg gccagaacca gctctataac	1200
gagctcaatc taggacgaag agaggagtac gatgttttgg acaagagacg tggccgggac	1260
cctgagatgg ggggaaagcc gagaaggaag aaccctcagg aaggcctgtt caatgaactg	1320
cagaaagata agatggcgga ggccttcagt gagattggga tgaaaggcga gcgccggagg	1380
ggcaaggggc acgatggcct tttccagggt ctcatgacag ccaccaagga caccttcgac	1440
gcccttcaca tgcaggccct gcccctcgc tgataa	1476

<210> 1595

<211> 490

<212> PRT

<213> Artificial Sequence

<220>

<223> mu20A10CAR T8283z1XX

<400> 1595

Met	Ala	Leu	Pro	Val	Thr	Ala	Leu	Leu	Leu	Pro	Leu	Ala	Leu	Leu	Leu
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His Ala Ala Arg Pro Glu Val Met Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Thr Tyr Ala Met Ser Trp Ile Arg Gln Thr Pro Glu Lys  
50 55 60

Arg Leu Glu Trp Val Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr  
65 70 75 80

Ser Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Val Arg  
85 90 95

Asn Ile Leu Tyr Leu Gln Met Ser Ser Leu Arg Ser Glu Asp Thr Ala  
100 105 110

Met Tyr Tyr Cys Ala Arg Gly Pro Ile Tyr Asn Asp Tyr Asp Glu Phe  
115 120 125

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala Gly Gly Gly  
130 135 140

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asn Ile Met Met  
145 150 155 160

Thr Gln Ser Pro Ser Ser Leu Ala Val Ser Ala Gly Glu Lys Val Thr  
165 170 175

Met Ser Cys Lys Ser Ser Gln Ser Val Leu Tyr Ser Ser Asn Gln Lys  
180 185 190

Asn Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ser Pro Lys Leu  
195 200 205

Leu Ile Tyr Trp Ala Ser Thr Arg Glu Ser Gly Val Pro Asp Arg Phe  
210 215 220

Thr Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Val  
225 230 235 240

Gln Ala Glu Asp Leu Ala Val Tyr Tyr Cys His Gln Tyr Leu Ser Ser  
245 250 255

Leu Thr Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys Thr Thr Thr Pro  
260 265 270

Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu  
275 280 285

Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His

290

295

300

Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu  
305 310 315 320

Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr  
325 330 335

Cys Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met  
340 345 350

Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala  
355 360 365

Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser Arg  
370 375 380

Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn  
385 390 395 400

Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg  
405 410 415

Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro  
420 425 430

Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala  
435 440 445

Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His  
450 455 460

Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe Asp  
465 470 475 480

Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

&lt;210&gt; 1596

&lt;211&gt; 1479

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; hu20A10CAR T8283z1XX

&lt;400&gt; 1596

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ccggagggtgc agctggttga atctggcggc ggacttgtga agcctggcgg atctctgaga 120

ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg ggtccgacag 180

gcccttgaa aaggccttga atgggttgcc tctatcgga gagccggcag cacctactac 240

agcgattctg tgaagggcag attcaccatc agccgggaca acgccaagaa cagcctgtac 300

ctgcagatga actccctgag agccgaggac accgccgtgt actattgtgc cagaggaccc 360  
atctacaacg actacgacga gttcgcctat tggggccagg gcacactggt cacagtcagc 420  
tctggcggtg gcggaagcgg aggcggtggc tccggtggcg gaggcagcga cattgtgctg 480  
accagtctc cagcctcctt ggccgtgtct ccaggacaga gggccaccat cacctgcaag 540  
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag 600  
aaaccaggac aacctcctaa actcctgatt tactgggccg gcaccagaga aagcggggtc 660  
ccagccaggt tcagcggcag tgggtctggg accgatttca ccctcacaat taatcctgtg 720  
gaagctaatt atactgcaaa ttattactgt caccagtacc tgagcagcct gaccttcggc 780  
ggagggacca aggtggagat caaacgaaca acaaccctg ccccagacc tcctaccca 840  
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ggcggagccg tgcacaccag aggactggat ttgcctgcg acatctacat ctgggcgccc 960  
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gaccttgaga tggggggaaa gccgagaagg aagaaccctc aggaaggcct gttcaatgaa 1320  
ctgcagaaag ataagatggc ggaggccttc agtgagattg ggatgaaagg cgagcgccgg 1380  
aggggcaagg ggcacgatgg ctttttcag ggtctcagta cagccaccaa ggacaccttc 1440  
gacgcccttc acatgcaggc cctgccccct cgctgataa 1479

<210> 1597  
<211> 491  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> hu20A10CAR T8283z1XX

<400> 1597

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Thr Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Gly Leu Glu Trp Val Ala Ser Ile Gly Arg Ala Gly Ser Thr Tyr Tyr

65				70					75					80			
Ser	Asp	Ser	Val	Lys 85	Gly	Arg	Phe	Thr	Ile 90	Ser	Arg	Asp	Asn	Ala 95	Lys		
Asn	Ser	Leu	Tyr 100	Leu	Gln	Met	Asn	Ser 105	Leu	Arg	Ala	Glu	Asp 110	Thr	Ala		
Val	Tyr	Tyr 115	Cys	Ala	Arg	Gly	Pro 120	Ile	Tyr	Asn	Asp	Tyr 125	Asp	Glu	Phe		
Ala	Tyr	Trp	Gly	Gln	Gly	Thr 135	Leu	Val	Thr	Val	Ser 140	Ser	Gly	Gly	Gly		
Gly 145	Ser	Gly	Gly	Gly	Gly 150	Ser	Gly	Gly	Gly 155	Ser	Asp	Ile	Val	Leu 160			
Thr	Gln	Ser	Pro	Ala 165	Ser	Leu	Ala	Val	Ser 170	Pro	Gly	Gln	Arg	Ala 175	Thr		
Ile	Thr	Cys	Lys 180	Ser	Ser	Gln	Ser	Val 185	Leu	Tyr	Ser	Ser	Asn 190	Gln	Lys		
Asn	Tyr	Leu 195	Ala	Trp	Tyr	Gln	Gln 200	Lys	Pro	Gly	Gln 205	Pro	Pro	Lys	Leu		
Leu	Ile 210	Tyr	Trp	Ala	Ser	Thr 215	Arg	Glu	Ser	Gly	Val 220	Pro	Ala	Arg	Phe		
Ser 225	Gly	Ser	Gly	Ser	Gly 230	Thr	Asp	Phe	Thr	Leu 235	Thr	Ile	Asn	Pro	Val 240		
Glu	Ala	Asn	Asp	Thr 245	Ala	Asn	Tyr	Tyr	Cys 250	His	Gln	Tyr	Leu	Ser 255	Ser		
Leu	Thr	Phe	Gly 260	Gly	Gly	Thr	Lys	Val 265	Glu	Ile	Lys	Arg	Thr 270	Thr	Thr		
Pro	Ala	Pro 275	Arg	Pro	Pro	Thr	Pro 280	Ala	Pro	Thr	Ile	Ala 285	Ser	Gln	Pro		
Leu	Ser 290	Leu	Arg	Pro	Glu	Ala 295	Cys	Arg	Pro	Ala	Ala 300	Gly	Gly	Ala	Val		
His 305	Thr	Arg	Gly	Leu	Asp 310	Phe	Ala	Cys	Asp	Ile 315	Tyr	Ile	Trp	Ala	Pro 320		
Leu	Ala	Gly	Thr	Cys 325	Gly	Val	Leu	Leu	Leu 330	Ser	Leu	Val	Ile	Thr 335	Leu		
Tyr	Cys	Arg	Ser 340	Lys	Arg	Ser	Arg	Leu 345	Leu	His	Ser	Asp	Tyr 350	Met	Asn		



Met Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr  
355 360 365

Ala Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser  
370 375 380

Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr  
385 390 395 400

Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys  
405 410 415

Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn  
420 425 430

Pro Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu  
435 440 445

Ala Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly  
450 455 460

His Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe  
465 470 475 480

Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1598

<211> 738

<212> DNA

<213> Artificial Sequence

<220>

<223> mu25E6 scFv full sequence

<400> 1598

gaggtgcagc tggaggagtc tgggggagac ttagtgaagc ctggagggtc cctgaaactc 60

tcctgtgcag cctctggttt cactttcagt agttatggaa tgtcttggtg tcgccagact 120

ccagacaaga ggctggagtg ggtcgcaacc attagtaatg gtggtagaca caccttctat 180

ccagacagtg tgaaggggcg attcaccatc tccagagaca atgccaagaa caccctgtat 240

ctgcaaatga gcagtctgaa gtctgaggac acagccatgt atttatgtgt aagacagact 300

gggacggagg gctggtttgc ttactggggc caagggactc tggtcactgt ctctgcaggt 360

ggcggaggat ctggcggagg tgggaagcggc ggaggcggat ccgatgttgt gatgaccag 420

actccactca ctttgtcggg taccattgga caaccagcct ccatctcttg caagtcaagt 480

cagagcctct tagatagtga tggaaagaca tatttgaatt ggttggtaca gaggccaggc 540

cagtctccaa agcgcctaata ctatctgggt tctaaactgg actctggagt ccctgacagg 600

ttcactggca gtggatcagg gacagatttc acactgaaaa tcagcagagt ggaggctgag 660

gatttgggag tttattattg ctggcaaggt acacattttc ctcagacgtt cggtggaggc 720  
 accaagctgg aaatcaaa 738

<210> 1599  
 <211> 246  
 <212> PRT  
 <213> Artificial Sequence  
 <220>  
 <223> mu25E6 scFv full sequence

<400> 1599

Glu Val Gln Leu Val Glu Ser Gly Gly Asp Leu Val Lys Pro Gly Gly  
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Ser Tyr  
 20 25 30

Gly Met Ser Trp Val Arg Gln Thr Pro Asp Lys Arg Leu Glu Trp Val  
 35 40 45

Ala Thr Ile Ser Asn Gly Gly Arg His Thr Phe Tyr Pro Asp Ser Val  
 50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Thr Leu Tyr  
 65 70 75 80

Leu Gln Met Ser Ser Leu Lys Ser Glu Asp Thr Ala Met Tyr Leu Cys  
 85 90 95

Val Arg Gln Thr Gly Thr Glu Gly Trp Phe Ala Tyr Trp Gly Gln Gly  
 100 105 110

Thr Leu Val Thr Val Ser Ala Gly Gly Gly Gly Ser Gly Gly Gly Gly  
 115 120 125

Ser Gly Gly Gly Gly Ser Asp Val Val Met Thr Gln Thr Pro Leu Thr  
 130 135 140

Leu Ser Val Thr Ile Gly Gln Pro Ala Ser Ile Ser Cys Lys Ser Ser  
 145 150 155 160

Gln Ser Leu Leu Asp Ser Asp Gly Lys Thr Tyr Leu Asn Trp Leu Leu  
 165 170 175

Gln Arg Pro Gly Gln Ser Pro Lys Arg Leu Ile Tyr Leu Val Ser Lys  
 180 185 190

Leu Asp Ser Gly Val Pro Asp Arg Phe Thr Gly Ser Gly Ser Gly Thr  
 195 200 205

Asp Phe Thr Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Leu Gly Val  
 210 215 220

Tyr Tyr Cys Trp Gln Gly Thr His Phe Pro Gln Thr Phe Gly Gly Gly  
225 230 235 240

Thr Lys Leu Glu Ile Lys  
245

<210> 1600  
<211> 738  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> hu25E6 full sequence

<400> 1600  
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cctggcaagg gcctggaatg ggtcagcacc atcagcaacg gcggaagaca caccttctac 180  
cccgacagcg tgaagggcag attcaccatc tcaagagata acgccaagaa cagcctgtac 240  
ctgcagatga acagcctgcg ggccgaggac accgccgtgt actactgcgc cagacagacc 300  
ggcacagagg gctggttcgc ctactggggc cagggcaccc tggtgaccgt gtccagcggc 360  
ggtggcggaa gcggaggcgg tggctccggt ggccgaggca gcgacatcgt gatgaccag 420  
acccctctgt ctctgagcgt gaccctggc cagcctgcca gcatctcttg taaaagcagc 480  
cagagcctgc tggacagcga cggcaagacc tacctgaact ggtacctgca gaagcccggc 540  
caaagccctc agctgctgat ctacctggtg tccaagctgg atagcggtgt tcctgataga 600  
ttcagcggat ctggcagcgg caccgacttc accctgaaga tcagcagagt ggaagccgag 660  
gacgtgggcg tgtactactg ctggcagggc acacacttcc cccagacatt cggccagggc 720  
accaaggtgg aaatcaag 738

<210> 1601  
<211> 246  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> hu25E6 full sequence

<400> 1601

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Lys Pro Gly Gly  
1 5 10 15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Ser Tyr  
20 25 30

Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val  
35 40 45

Ser Thr Ile Ser Asn Gly Gly Arg His Thr Phe Tyr Pro Asp Ser Val

50

55

60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr  
65 70 75 80

Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys  
85 90 95

Ala Arg Gln Thr Gly Thr Glu Gly Trp Phe Ala Tyr Trp Gly Gln Gly  
100 105 110

Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly  
115 120 125

Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Thr Pro Leu Ser  
130 135 140

Leu Ser Val Thr Pro Gly Gln Pro Ala Ser Ile Ser Cys Lys Ser Ser  
145 150 155 160

Gln Ser Leu Leu Asp Ser Asp Gly Lys Thr Tyr Leu Asn Trp Tyr Leu  
165 170 175

Gln Lys Pro Gly Gln Ser Pro Gln Leu Leu Ile Tyr Leu Val Ser Lys  
180 185 190

Leu Asp Ser Gly Val Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr  
195 200 205

Asp Phe Thr Leu Lys Ile Ser Arg Val Glu Ala Glu Asp Val Gly Val  
210 215 220

Tyr Tyr Cys Trp Gln Gly Thr His Phe Pro Gln Thr Phe Gly Gln Gly  
225 230 235 240

Thr Lys Val Glu Ile Lys  
245

<210> 1602

<211> 1476

<212> DNA

<213> Artificial Sequence

<220>

<223> mu25E6CAR T841BB3z

<400> 1602

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ctctcctgtg cagcctctgg ttctactttc agtagttatg gaatgtcttg ggttcgccag 180

actccagaca agaggctgga gtgggtcgca accattagta atgggtgtag acacaccttc 240

tatccagaca gtgtgaaggg gcgattcacc atctccagag acaatgcaa gaacaccctg 300

tatctgcaaa tgagcagtct gaagtctgag gacacagcca tgtatttatg tgtaagacag 360  
actgggacgg agggctgggt tgcttactgg ggccaaggga ctctgggtcac tgtctctgca 420  
ggtagcgagg gatctggcgg aggtggaagc ggaggaggcg gatccgatgt tgtgatgacc 480  
cagactccac tcactttgtc ggttaccatt ggacaaccag cctccatctc ttgcaagtca 540  
agtcagagcc tcttagatag tgatggaaag acatatattga attggttgtt acagaggcca 600  
ggccagtctc caaagcgctt aatctatctg gtgtctaaac tggactctgg agtccctgac 660  
aggttcactg gcagtggatc agggacagat ttcacactga aaatcagcag agtggaggct 720  
gaggatttgg gaggtttatta ttgctggcaa ggtacacatt ttcctcagac gttcgggtgga 780  
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acaattgccg gccagcctct gagcctgagg cccgaggcct gtagacctgc tgctggcgga 900  
gccgtgcaca ccagaggact ggatttcgcc tgcgacatct acatctgggc gcccttggcc 960  
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aagaaactcc tgtatatatt caaacaacca tttatgagac cagtacaaac tactcaagag 1080  
gaagatggct gtagctgccg atttccagaa gaagaagaag gaggatgtga actgagagtg 1140  
aagttcagca ggagcgcaga cggccccgcg tacaagcagg gccagaacca gctctataac 1200  
gagctcaatc taggacgaag agaggagtac gatgttttgg acaagagacg tggccgggac 1260  
cctgagatgg ggggaaagcc gagaaggaag aaccctcagg aaggcctgta caatgaactg 1320  
cagaaagata agatggcgga ggacctacgt gagattggga tgaaaggcga gcgccggagg 1380  
ggcaaggggc acgatggcct ttaccagggt ctcagtacag ccaccaagga cacctacgac 1440  
gcccttcaca tgcaggccct gccccctcgc tgataa 1476

<210> 1603  
<211> 490  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> mu25E6CAR T841BB3z

<400> 1603

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Asp Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Ser Tyr Gly Met Ser Trp Val Arg Gln Thr Pro Asp Lys  
50 55 60

Arg Leu Glu Trp Val Ala Thr Ile Ser Asn Gly Gly Arg His Thr Phe

65					70						75				80
Tyr	Pro	Asp	Ser	Val	Lys	Gly	Arg	Phe	Thr	Ile	Ser	Arg	Asp	Asn	Ala
			85						90					95	
Lys	Asn	Thr	Leu	Tyr	Leu	Gln	Met	Ser	Ser	Leu	Lys	Ser	Glu	Asp	Thr
			100					105					110		
Ala	Met	Tyr	Leu	Cys	Val	Arg	Gln	Thr	Gly	Thr	Glu	Gly	Trp	Phe	Ala
		115					120					125			
Tyr	Trp	Gly	Gln	Gly	Thr	Leu	Val	Thr	Val	Ser	Ala	Gly	Gly	Gly	Gly
	130					135					140				
Ser	Gly	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser	Asp	Val	Val	Met	Thr
145					150					155					160
Gln	Thr	Pro	Leu	Thr	Leu	Ser	Val	Thr	Ile	Gly	Gln	Pro	Ala	Ser	Ile
			165						170					175	
Ser	Cys	Lys	Ser	Ser	Gln	Ser	Leu	Leu	Asp	Ser	Asp	Gly	Lys	Thr	Tyr
			180					185					190		
Leu	Asn	Trp	Leu	Leu	Gln	Arg	Pro	Gly	Gln	Ser	Pro	Lys	Arg	Leu	Ile
		195					200					205			
Tyr	Leu	Val	Ser	Lys	Leu	Asp	Ser	Gly	Val	Pro	Asp	Arg	Phe	Thr	Gly
	210					215					220				
Ser	Gly	Ser	Gly	Thr	Asp	Phe	Thr	Leu	Lys	Ile	Ser	Arg	Val	Glu	Ala
225					230					235					240
Glu	Asp	Leu	Gly	Val	Tyr	Tyr	Cys	Trp	Gln	Gly	Thr	His	Phe	Pro	Gln
				245					250					255	
Thr	Phe	Gly	Gly	Gly	Thr	Lys	Leu	Glu	Ile	Lys	Thr	Thr	Thr	Pro	Ala
		260						265						270	
Pro	Arg	Pro	Pro	Thr	Pro	Ala	Pro	Thr	Ile	Ala	Ser	Gln	Pro	Leu	Ser
		275					280					285			
Leu	Arg	Pro	Glu	Ala	Cys	Arg	Pro	Ala	Ala	Gly	Gly	Ala	Val	His	Thr
	290					295					300				
Arg	Gly	Leu	Asp	Phe	Ala	Cys	Asp	Ile	Tyr	Ile	Trp	Ala	Pro	Leu	Ala
305					310					315					320
Gly	Thr	Cys	Gly	Val	Leu	Leu	Leu	Ser	Leu	Val	Ile	Thr	Leu	Tyr	Cys
				325					330					335	
Lys	Arg	Gly	Arg	Lys	Lys	Leu	Leu	Tyr	Ile	Phe	Lys	Gln	Pro	Phe	Met
			340					345					350		

Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe  
355 360 365

Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser Arg  
370 375 380

Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn  
385 390 395 400

Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg  
405 410 415

Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro  
420 425 430

Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala  
435 440 445

Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His  
450 455 460

Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr Asp  
465 470 475 480

Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1604  
<211> 1476  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> hu25E6CAR T841BB3z

<400> 1604  
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ctgagctgcg ccgccagcgg cttcacattc agcagctacg gcatgagctg ggtgcggcag 180  
gcccctggca agggcctgga atgggtcagc accatcagca acggcggaag acacaccttc 240  
taccgccgaca gcgtgaaggg cagattcacc atctcaagag ataacgcaa gaacagcctg 300  
tacctgcaga tgaacagcct gcgggccgag gacaccgccg tgtactactg cgccagacag 360  
accggcacag agggctgggtt cgcctactgg ggccagggca ccctgggtgac cgtgtccagc 420  
ggcgggtggcg gaagcggagg cgggtggctcc ggtggcggag gcagcgacat cgtgatgacc 480  
cagacccttc tgtctctgag cgtgaccctt ggccagcctg ccagcatctc ttgtaaaagc 540  
agccagagcc tgctggacag cgacggcaag acctacctga actggtacct gcagaagccc 600  
ggccaaagcc ctcagctgct gatctacctg gtgtccaagc tggatagcgg tgttcctgat 660

agattcagcg gatctggcag cggcaccgac ttcaccctga agatcagcag agtggaagcc 720  
gaggacgtgg gcgtgtacta ctgctggcag ggcacacact tccccagac attcggccag 780  
ggcaccaagg tggaaatcaa gacaacaacc cctgccccca gacctctac cccagcccct 840  
acaattgcc a gccagcctct gagcctgagg cccgaggcct gtagacctgc tgctggcgga 900  
gccgtgcaca ccagaggact ggatttcgcc tgcgacatct acatctgggc gcccttggcc 960  
gggacttggt gggtccttct cctgtcactg gttatcaccc tttactgcaa acggggcaga 1020  
aagaaactcc tgtatatatt caaacaacca tttatgagac cagtacaaac tactcaagag 1080  
gaagatggct gtagctgccg atttccagaa gaagaagaag gaggatgtga actgagagtg 1140  
aagttcagca ggagcgcaga cggccccgcg tacaagcagg gccagaacca gctctataac 1200  
gagctcaatc taggacgaag agaggagtac gatgttttgg acaagagacg tggccgggac 1260  
cctgagatgg ggggaaagcc gagaaggaag aaccctcagg aaggcctgta caatgaactg 1320  
cagaaagata agatggcgga ggcctacagt gagattggga tgaaaggcga gcgccggagg 1380  
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gcccttcaca tgcaggccct gccccctcgc tgataa 1476

<210> 1605  
<211> 490  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> hu25E6CAR T841BB3z

<400> 1605

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Ser Tyr Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Gly Leu Glu Trp Val Ser Thr Ile Ser Asn Gly Gly Arg His Thr Phe  
65 70 75 80

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95

Lys Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110

Ala Val Tyr Tyr Cys Ala Arg Gln Thr Gly Thr Glu Gly Trp Phe Ala  
115 120 125



Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly  
 130 135 140

Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr  
 145 150 155 160

Gln Thr Pro Leu Ser Leu Ser Val Thr Pro Gly Gln Pro Ala Ser Ile  
 165 170 175

Ser Cys Lys Ser Ser Gln Ser Leu Leu Asp Ser Asp Gly Lys Thr Tyr  
 180 185 190

Leu Asn Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Gln Leu Leu Ile  
 195 200 205

Tyr Leu Val Ser Lys Leu Asp Ser Gly Val Pro Asp Arg Phe Ser Gly  
 210 215 220

Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile Ser Arg Val Glu Ala  
 225 230 235 240

Glu Asp Val Gly Val Tyr Tyr Cys Trp Gln Gly Thr His Phe Pro Gln  
 245 250 255

Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys Thr Thr Thr Pro Ala  
 260 265 270

Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser  
 275 280 285

Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr  
 290 295 300

Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala  
 305 310 315 320

Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys  
 325 330 335

Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe Met  
 340 345 350

Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe  
 355 360 365

Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser Arg  
 370 375 380

Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn  
 385 390 395 400

Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg  
405 410 415

Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro  
420 425 430

Gln Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala  
435 440 445

Tyr Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His  
450 455 460

Asp Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr Asp  
465 470 475 480

Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1606  
<211> 1473  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> mu25E6CAR T8283z

<400> 1606  
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ctctcctgtg cagcctctgg tttcactttc agtagttatg gaatgtcttg ggttcgccag 180  
actccagaca agaggctgga gtgggtcgca accattagta atgggtggtag acacaccttc 240  
tatccagaca gtgtgaaggg gcgattcacc atctccagag acaatgcaa gaacaccctg 300  
tatctgcaaa tgagcagtct gaagtctgag gacacagcca tgtatttatg tgtaagacag 360  
actgggacgg agggctgggtt tgcttactgg ggccaaggga ctctgggtcac tgtctctgca 420  
ggtggcggag gatctggcgg aggtggaagc ggcggaggcg gatccgatgt tgtgatgacc 480  
cagactccac tcactttgtc ggttaccatt ggacaaccag cctccatctc ttgcaagtca 540  
agtcagagcc tcttagatag tgatggaaa acatatttga attggttggt acagaggcca 600  
ggccagtctc caaagcgctt aatctatctg gtgtctaaac tggactctgg agtccctgac 660  
aggttcactg gcagtggatc agggacagat ttcacactga aaatcagcag agtggaggct 720  
gaggatttgg gattttatta ttgctggcaa ggtacacatt ttcctcagac gttcgggtgga 780  
ggcaccaagc tggaaatcaa aacaacaacc cctgccccca gacctcctac cccagcccct 840  
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gccgtgcaca ccagaggact ggatttcgcc tgcgacatct acatctgggc gcccttggcc 960  
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aaagataaga tggcggaggc ctacagttag attgggatga aaggcgagcg ccggaggggc 1380  
aaggggcacg atggccttta ccagggtctc agtacagcca ccaaggacac ctacgacgcc 1440  
cttcacatgc aggccttgcc ccctcgctga taa 1473

<210> 1607  
<211> 489  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> mu25E6CAR T8283z

<400> 1607

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
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His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Asp Leu  
20 25 30  
  
Val Lys Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45  
  
Thr Phe Ser Ser Tyr Gly Met Ser Trp Val Arg Gln Thr Pro Asp Lys  
50 55 60  
  
Arg Leu Glu Trp Val Ala Thr Ile Ser Asn Gly Gly Arg His Thr Phe  
65 70 75 80  
  
Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95  
  
Lys Asn Thr Leu Tyr Leu Gln Met Ser Ser Leu Lys Ser Glu Asp Thr  
100 105 110  
  
Ala Met Tyr Leu Cys Val Arg Gln Thr Gly Thr Glu Gly Trp Phe Ala  
115 120 125  
  
Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala Gly Gly Gly Gly  
130 135 140  
  
Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Val Val Met Thr  
145 150 155 160  
  
Gln Thr Pro Leu Thr Leu Ser Val Thr Ile Gly Gln Pro Ala Ser Ile  
165 170 175

Ser Cys Lys Ser Ser Gln Ser Leu Leu Asp Ser Asp Gly Lys Thr Tyr  
 180 185 190

Leu Asn Trp Leu Leu Gln Arg Pro Gly Gln Ser Pro Lys Arg Leu Ile  
 195 200 205

Tyr Leu Val Ser Lys Leu Asp Ser Gly Val Pro Asp Arg Phe Thr Gly  
 210 215 220

Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile Ser Arg Val Glu Ala  
 225 230 235 240

Glu Asp Leu Gly Val Tyr Tyr Cys Trp Gln Gly Thr His Phe Pro Gln  
 245 250 255

Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys Thr Thr Thr Pro Ala  
 260 265 270

Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser  
 275 280 285

Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr  
 290 295 300

Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala  
 305 310 315 320

Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys  
 325 330 335

Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met Thr  
 340 345 350

Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala Pro  
 355 360 365

Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser Arg Ser  
 370 375 380

Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu  
 385 390 395 400

Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg  
 405 410 415

Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln  
 420 425 430

Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Tyr  
 435 440 445

Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp  
450 455 460

Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr Asp Ala  
465 470 475 480

Leu His Met Gln Ala Leu Pro Pro Arg  
485

<210> 1608  
<211> 1473  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> hu25E6CAR T8283z

<400> 1608  
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gcccctggca agggcctgga atgggtcagc accatcagca acggcggaag acacaccttc 240  
taccgccaca gcgtgaaggg cagattcacc atctcaagag ataacgcaa gaacagcctg 300  
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ggcaccaagg tggaaatcaa gacaacaacc cctgccccca gacctctac ccagcccct 840  
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aagcattacc agccctatgc cccaccacgc gacttcgcag cctatcgctc cagagtgaag 1140  
ttcagcagga gcgcagacgc ccccgctac aagcagggcc agaaccagct ctataacgag 1200  
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gagatggggg gaaagccgag aaggaagaac cctcaggaag gcctgtacaa tgaactgcag 1320  
aaagataaga tggcggaggc ctacagttag attgggatga aaggcgagcg ccggaggggc 1380  
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&lt;210&gt; 1609

&lt;211&gt; 489

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; hu25E6CAR T8283z

&lt;400&gt; 1609

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Ser Tyr Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Gly Leu Glu Trp Val Ser Thr Ile Ser Asn Gly Gly Arg His Thr Phe  
65 70 75 80

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95

Lys Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110

Ala Val Tyr Tyr Cys Ala Arg Gln Thr Gly Thr Glu Gly Trp Phe Ala  
115 120 125

Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly  
130 135 140

Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr  
145 150 155 160

Gln Thr Pro Leu Ser Leu Ser Val Thr Pro Gly Gln Pro Ala Ser Ile  
165 170 175

Ser Cys Lys Ser Ser Gln Ser Leu Leu Asp Ser Asp Gly Lys Thr Tyr  
180 185 190

Leu Asn Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Gln Leu Leu Ile  
195 200 205

Tyr Leu Val Ser Lys Leu Asp Ser Gly Val Pro Asp Arg Phe Ser Gly  
210 215 220

Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile Ser Arg Val Glu Ala  
225 230 235 240

Glu Asp Val Gly Val Tyr Tyr Cys Trp Gln Gly Thr His Phe Pro Gln  
245 250 255

Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys Thr Thr Thr Pro Ala  
260 265 270

Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser  
275 280 285

Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr  
290 295 300

Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala  
305 310 315 320

Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys  
325 330 335

Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met Thr  
340 345 350

Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala Pro  
355 360 365

Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser Arg Ser  
370 375 380

Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu  
385 390 395 400

Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg  
405 410 415

Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln  
420 425 430

Glu Gly Leu Tyr Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Tyr  
435 440 445

Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp  
450 455 460

Gly Leu Tyr Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Tyr Asp Ala  
465 470 475 480

Leu His Met Gln Ala Leu Pro Pro Arg  
485

<210> 1610

<211> 1476  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> mu25E6CAR T841BB3z1XX

<400> 1610  
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ctctcctgtg cagcctctgg tttcactttc agtagttatg gaatgtcttg ggttcgccag 180  
actccagaca agaggctgga gtgggtcgca accattagta atgggtgtag acacaccttc 240  
tatccagaca gtgtgaaggg gcgattcacc atctccagag acaatgcaa gaacaccctg 300  
tatctgcaaa tgagcagtct gaagtctgag gacacagcca tgtatttatg tgtaagacag 360  
actgggacgg agggctggtt tgcttactgg ggccaaggga ctctggtcac tgtctctgca 420  
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cagactccac tcactttgtc ggttaccatt ggacaaccag cctccatctc ttgcaagtca 540  
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aggttcactg gcagtggatc agggacagat ttcacactga aaatcagcag agtggaggct 720  
gaggatttgg gaggtttatta ttgctggcaa ggtacacatt ttcctcagac gttcggtgga 780  
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acaattgcca gccagcctct gagcctgagg cccgaggcct gtagacctgc tgctggcgga 900  
gccgtgcaca ccagaggact ggatttcgcc tgcgacatct acatctgggc gcccttggcc 960  
gggacttggt gggtccttct cctgtcactg gttatcacc tttactgcaa acggggcaga 1020  
aagaaactcc tgtatatatt caaacaacca tttatgagac cagtacaaac tactcaagag 1080  
gaagatggct gtagctgccg atttccagaa gaagaagaag gaggatgtga actgagagtg 1140  
aagttcagca ggagcgcaga cggccccgcg tacaagcagg gccagaacca gctctataac 1200  
gagctcaatc taggacgaag agaggagtac gatgttttgg acaagagacg tggccgggac 1260  
cctgagatgg ggggaaagcc gagaaggaag aaccctcagg aaggcctggt caatgaactg 1320  
cagaaagata agatggcgga ggccttcagt gagattggga tgaaaggcga gcgccggagg 1380  
ggcaaggggc acgatggcct tttccagggt ctcagtacag ccaccaagga caccttcgac 1440  
gcccttcaca tgcaggccct gccccctcgc tgataa 1476

<210> 1611  
<211> 490  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> mu25E6CAR T841BB3z1XX

<400> 1611



Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Asp Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Ser Tyr Gly Met Ser Trp Val Arg Gln Thr Pro Asp Lys  
50 55 60

Arg Leu Glu Trp Val Ala Thr Ile Ser Asn Gly Gly Arg His Thr Phe  
65 70 75 80

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95

Lys Asn Thr Leu Tyr Leu Gln Met Ser Ser Leu Lys Ser Glu Asp Thr  
100 105 110

Ala Met Tyr Leu Cys Val Arg Gln Thr Gly Thr Glu Gly Trp Phe Ala  
115 120 125

Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala Gly Gly Gly Gly  
130 135 140

Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Val Val Met Thr  
145 150 155 160

Gln Thr Pro Leu Thr Leu Ser Val Thr Ile Gly Gln Pro Ala Ser Ile  
165 170 175

Ser Cys Lys Ser Ser Gln Ser Leu Leu Asp Ser Asp Gly Lys Thr Tyr  
180 185 190

Leu Asn Trp Leu Leu Gln Arg Pro Gly Gln Ser Pro Lys Arg Leu Ile  
195 200 205

Tyr Leu Val Ser Lys Leu Asp Ser Gly Val Pro Asp Arg Phe Thr Gly  
210 215 220

Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile Ser Arg Val Glu Ala  
225 230 235 240

Glu Asp Leu Gly Val Tyr Tyr Cys Trp Gln Gly Thr His Phe Pro Gln  
245 250 255

Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys Thr Thr Thr Pro Ala  
260 265 270

Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser

275

280

285

Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr  
 290 295 300

Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala  
 305 310 315 320

Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys  
 325 330 335

Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe Met  
 340 345 350

Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe  
 355 360 365

Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser Arg  
 370 375 380

Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn  
 385 390 395 400

Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg  
 405 410 415

Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro  
 420 425 430

Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala  
 435 440 445

Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His  
 450 455 460

Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe Asp  
 465 470 475 480

Ala Leu His Met Gln Ala Leu Pro Pro Arg  
 485 490

&lt;210&gt; 1612

&lt;211&gt; 1476

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; hu25E6CAR T841BB3z1XX

&lt;400&gt; 1612

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ctgagctgcg ccgccagcgg cttcacattc agcagctacg gcatgagctg ggtgcggcag 180

gcccctggca agggcctgga atgggtcagc accatcagca acggcggaag acacaccttc 240  
taccccgaca gcgtgaaggg cagattcacc atctcaagag ataacgcaa gaacagcctg 300  
tacctgcaga tgaacagcct gcgggcccag gacaccgccg tgtactactg cgccagacag 360  
accggcacag agggctgggt cgcctactgg ggccagggca ccctgggtgac cgtgtccagc 420  
ggcgggtggcg gaagcggagg cgggtggctcc ggtggcggag gcagcgacat cgtgatgacc 480  
cagacccctc tgtctctgag cgtgaccctt ggccagcctg ccagcatctc ttgtaaaagc 540  
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ggccaaagcc ctcagctgct gatctacctg gtgtccaagc tggatagcgg tgttcctgat 660  
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gaggacgtgg gcgtgtacta ctgctggcag ggcacacact tccccagac attcggccag 780  
ggcaccaagg tggaaatcaa gacaacaacc cctgccccca gacctctac ccagcccct 840  
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gccgtgcaca ccagaggact ggatttcgcc tgcgacatct acatctgggc gcccttggcc 960  
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gaagatggct gtagctgccg atttccagaa gaagaagaag gaggatgtga actgagagtg 1140  
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gagctcaatc taggacgaag agaggagtac gatgttttgg acaagagacg tggccgggac 1260  
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cagaaagata agatggcgga ggccttcagt gagattggga tgaaaggcga gcgccggagg 1380  
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gcccttcaca tgcaggccct gccccctcgc tgataa 1476

<210> 1613  
<211> 490  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> hu25E6CAR T841BB3z1XX  
  
<400> 1613

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15  
  
His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30  
  
Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45  
  
Thr Phe Ser Ser Tyr Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys

50	55	60														
Gly 65	Leu	Glu	Trp	Val	Ser 70	Thr	Ile	Ser	Asn	Gly 75	Gly	Arg	His	Thr	Phe 80	
Tyr	Pro	Asp	Ser	Val 85	Lys	Gly	Arg	Phe	Thr 90	Ile	Ser	Arg	Asp	Asn 95	Ala	
Lys	Asn	Ser	Leu 100	Tyr	Leu	Gln	Met	Asn 105	Ser	Leu	Arg	Ala	Glu 110	Asp	Thr	
Ala	Val	Tyr	Tyr	Cys	Ala	Arg	Gln 120	Thr	Gly	Thr	Glu	Gly 125	Trp	Phe	Ala	
Tyr	Trp	Gly	Gln	Gly	Thr	Leu 135	Val	Thr	Val	Ser	Ser	Gly	Gly	Gly	Gly	
Ser 145	Gly	Gly	Gly	Gly	Ser 150	Gly	Gly	Gly	Gly	Ser 155	Asp	Ile	Val	Met	Thr 160	
Gln	Thr	Pro	Leu	Ser 165	Leu	Ser	Val	Thr	Pro 170	Gly	Gln	Pro	Ala	Ser 175	Ile	
Ser	Cys	Lys	Ser 180	Ser	Gln	Ser	Leu	Leu 185	Asp	Ser	Asp	Gly	Lys 190	Thr	Tyr	
Leu	Asn	Trp	Tyr	Leu	Gln	Lys	Pro 200	Gly	Gln	Ser	Pro	Gln 205	Leu	Leu	Ile	
Tyr	Leu	Val	Ser	Lys	Leu	Asp 215	Ser	Gly	Val	Pro	Asp 220	Arg	Phe	Ser	Gly	
Ser 225	Gly	Ser	Gly	Thr	Asp 230	Phe	Thr	Leu	Lys	Ile 235	Ser	Arg	Val	Glu	Ala 240	
Glu	Asp	Val	Gly	Val 245	Tyr	Tyr	Cys	Trp	Gln 250	Gly	Thr	His	Phe	Pro 255	Gln	
Thr	Phe	Gly	Gln 260	Gly	Thr	Lys	Val	Glu 265	Ile	Lys	Thr	Thr	Thr 270	Pro	Ala	
Pro	Arg	Pro	Pro	Thr	Pro	Ala	Pro 280	Thr	Ile	Ala	Ser	Gln 285	Pro	Leu	Ser	
Leu	Arg 290	Pro	Glu	Ala	Cys	Arg 295	Pro	Ala	Ala	Gly	Gly 300	Ala	Val	His	Thr	
Arg 305	Gly	Leu	Asp	Phe	Ala 310	Cys	Asp	Ile	Tyr	Ile 315	Trp	Ala	Pro	Leu	Ala 320	
Gly	Thr	Cys	Gly	Val 325	Leu	Leu	Leu	Ser	Leu 330	Val	Ile	Thr	Leu	Tyr 335	Cys	

Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe Met  
340 345 350

Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe  
355 360 365

Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser Arg  
370 375 380

Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn  
385 390 395 400

Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg  
405 410 415

Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro  
420 425 430

Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala  
435 440 445

Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His  
450 455 460

Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe Asp  
465 470 475 480

Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1614  
<211> 1473  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> mu25E6CAR T8283z1XX

<400> 1614  
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tatccagaca gtgtgaaggg gcgattcacc atctccagag acaatgcaa gaacaccctg 300  
tatctgcaaa tgagcagtct gaagtctgag gacacagcca tgtatttatg tgtaagacag 360  
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gggtggcggag gatctggcgg aggtggaagc ggcggaggcg gatccgatgt tgtgatgacc 480  
cagactccac tcactttgtc gggttaccatt ggacaaccag cctccatctc ttgcaagtca 540

agtcagagcc tcttagatag tgatggaaag acatatattga attggttggt acagaggcca 600  
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gaggatttgg gattttatta ttgctggcaa ggtacacatt ttcctcagac gttcgggtgga 780  
ggcaccaagc tggaaatcaa aacaacaacc cctgccccca gacctcctac cccagcccct 840  
acaattgcc a gccagcctct gagcctgagg cccgaggctt gtagacctgc tgctggcgga 900  
gccgtgcaca ccagaggact ggatttcgcc tgcgacatct acatctgggc gcccttggcc 960  
gggacttgtg gggtccttct cctgtcactg gttatcaccc tttactgcag gagtaagagg 1020  
agcaggctcc tgcacagtga ctacatgaac atgactccta gaagacctgg gcctaccaga 1080  
aagcattacc agccctatgc cccaccacgc gacttcgcag cctatcgctc cagagtgaag 1140  
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ctcaatctag gacgaagaga ggagtacgat gttttggaca agagacgtgg ccgggaccct 1260  
gagatggggg gaaagccgag aaggaagaac cctcaggaag gcctgttcaa tgaactgcag 1320  
aaagataaga tggcggaggc cttcagtgcg attgggatga aaggcgagcg ccggaggggc 1380  
aaggggcacg atggcctttt ccagggtctc agtacagcca ccaaggacac cttcgacgcc 1440  
cttcacatgc aggccctgcc ccctcgctga taa 1473

<210> 1615  
<211> 489  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> mu25E6CAR T8283z1XX  
  
<400> 1615

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15  
  
His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Asp Leu  
20 25 30  
  
Val Lys Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45  
  
Thr Phe Ser Ser Tyr Gly Met Ser Trp Val Arg Gln Thr Pro Asp Lys  
50 55 60  
  
Arg Leu Glu Trp Val Ala Thr Ile Ser Asn Gly Gly Arg His Thr Phe  
65 70 75 80  
  
Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95  
  
Lys Asn Thr Leu Tyr Leu Gln Met Ser Ser Leu Lys Ser Glu Asp Thr  
100 105 110

Ala Met Tyr Leu Cys Val Arg Gln Thr Gly Thr Glu Gly Trp Phe Ala  
 115 120 125

Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala Gly Gly Gly Gly  
 130 135 140

Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Val Val Met Thr  
 145 150 155 160

Gln Thr Pro Leu Thr Leu Ser Val Thr Ile Gly Gln Pro Ala Ser Ile  
 165 170 175

Ser Cys Lys Ser Ser Gln Ser Leu Leu Asp Ser Asp Gly Lys Thr Tyr  
 180 185 190

Leu Asn Trp Leu Leu Gln Arg Pro Gly Gln Ser Pro Lys Arg Leu Ile  
 195 200 205

Tyr Leu Val Ser Lys Leu Asp Ser Gly Val Pro Asp Arg Phe Thr Gly  
 210 215 220

Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile Ser Arg Val Glu Ala  
 225 230 235 240

Glu Asp Leu Gly Val Tyr Tyr Cys Trp Gln Gly Thr His Phe Pro Gln  
 245 250 255

Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys Thr Thr Thr Pro Ala  
 260 265 270

Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser  
 275 280 285

Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr  
 290 295 300

Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala  
 305 310 315 320

Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys  
 325 330 335

Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met Thr  
 340 345 350

Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala Pro  
 355 360 365

Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser Arg Ser  
 370 375 380

Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu  
385 390 395 400

Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg  
405 410 415

Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln  
420 425 430

Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Phe  
435 440 445

Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp  
450 455 460

Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe Asp Ala  
465 470 475 480

Leu His Met Gln Ala Leu Pro Pro Arg  
485

<210> 1616  
<211> 1473  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> hu25E6CAR T8283z1XX

<400> 1616  
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ccggagggtgc agctggtgga atctggcgga ggactggtca agcctggagg cagcctgaga 120  
ctgagctgcg ccgccagcgg cttcacattc agcagctacg gcatgagctg ggtgcggcag 180  
gcccctggca agggcctgga atgggtcagc accatcagca acggcggaag acacaccttc 240  
taccgccgaca gcgtgaaggg cagattcacc atctcaagag ataacgcaa gaacagcctg 300  
tacctgcaga tgaacagcct gcgggccgag gacaccgccg tgtactactg cgccagacag 360  
accggcacag agggctggtt cgcctactgg ggccagggca ccctggtgac cgtgtccagc 420  
ggcgggtggcg gaagcggagg cggtggctcc ggtggcggag gcagcgacat cgtgatgacc 480  
cagacccttc tgtctctgag cgtgaccctt ggccagcctg ccagcatctc ttgtaaaagc 540  
agccagagcc tgctggacag cgacggcaag acctacctga actggtacct gcagaagccc 600  
ggccaaagcc ctcagctgct gatctacctg gtgtccaagc tggatagcgg tgttcctgat 660  
agattcagcg gatctggcag cggcaccgac ttcaccctga agatcagcag agtggaagcc 720  
gaggacgtgg gcgtgtacta ctgctggcag ggcacacact tccccagac attcggccag 780  
ggcaccaagg tggaaatcaa gacaacaacc cctgccccca gacctctac ccagcccct 840  
acaattgcc a gccagcctct gagcctgagg ccgagggtt gtagacctgc tgctggcggga 900  
gccgtgcaca ccagaggact ggatttcgcc tgcgacatct acatctgggc gcccttggcc 960



gggacttgtg gggtccttct cctgtcactg gttatcaccc tttactgcag gagtaagagg 1020  
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aagcattacc agccctatgc cccaccacgc gacttcgcag cctatcgctc cagagtgaag 1140  
ttcagcagga gcgcagacgc ccccgctac aagcagggcc agaaccagct ctataacgag 1200  
ctcaatctag gacgaagaga ggagtacgat gttttggaca agagacgtgg ccgggaccct 1260  
gagatggggg gaaagccgag aaggaagaac cctcaggaag gcctgttcaa tgaactgcag 1320  
aaagataaga tggcggaggc cttcagttag attgggatga aaggcgagcg ccggaggggc 1380  
aaggggcacg atggcctttt ccagggtctc agtacagcca ccaaggacac cttcgacgcc 1440  
cttcacatgc aggccctgcc ccctcgctga taa 1473

<210> 1617  
<211> 489  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> hu25E6CAR T8283z1XX

<400> 1617

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15  
  
His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30  
  
Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45  
  
Thr Phe Ser Ser Tyr Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60  
  
Gly Leu Glu Trp Val Ser Thr Ile Ser Asn Gly Gly Arg His Thr Phe  
65 70 75 80  
  
Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95  
  
Lys Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110  
  
Ala Val Tyr Tyr Cys Ala Arg Gln Thr Gly Thr Glu Gly Trp Phe Ala  
115 120 125  
  
Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly  
130 135 140  
  
Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr  
145 150 155 160

Gln Thr Pro Leu Ser Leu Ser Val Thr Pro Gly Gln Pro Ala Ser Ile  
 165 170 175

Ser Cys Lys Ser Ser Gln Ser Leu Leu Asp Ser Asp Gly Lys Thr Tyr  
 180 185 190

Leu Asn Trp Tyr Leu Gln Lys Pro Gly Gln Ser Pro Gln Leu Leu Ile  
 195 200 205

Tyr Leu Val Ser Lys Leu Asp Ser Gly Val Pro Asp Arg Phe Ser Gly  
 210 215 220

Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile Ser Arg Val Glu Ala  
 225 230 235 240

Glu Asp Val Gly Val Tyr Tyr Cys Trp Gln Gly Thr His Phe Pro Gln  
 245 250 255

Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys Thr Thr Thr Pro Ala  
 260 265 270

Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser  
 275 280 285

Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr  
 290 295 300

Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala  
 305 310 315 320

Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys  
 325 330 335

Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met Thr  
 340 345 350

Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala Pro  
 355 360 365

Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser Arg Ser  
 370 375 380

Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu  
 385 390 395 400

Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg  
 405 410 415

Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln  
 420 425 430

Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Phe  
435 440 445

Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp  
450 455 460

Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe Asp Ala  
465 470 475 480

Leu His Met Gln Ala Leu Pro Pro Arg  
485

- <210> 1618
- <211> 1533
- <212> DNA
- <213> Artificial Sequence
- <220>
- <223> MNC21XX - muMNC2CAR T841BB3z1XX

<400> 1618  
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ccggagggtcc agctggagga gtcaggggga ggcttagtga agcctggagg gtccctgaaa 120  
ctctcctgtg cagcctctgg attcactttc agtggctatg ccatgtcttg ggttcgccag 180  
actccggaga agaggctgga gtgggtcgca accattagta gtggtggtac ttatatctac 240  
tatccagaca gtgtgaaggg gcgattcacc atctccagag acaatgcaa gaacaccctg 300  
tacctgcaaa tgagcagtct gaggtctgag gacacggcca tgtattactg tgcaagactt 360  
gggggggata attactacga atacttcgat gtctggggcg cagggaccac ggtcaccgtc 420  
tcctccgcca aaacgacacc cccatctgtc tatggcggtg gcggatccgg cgggtggcgga 480  
tccggcggtg gcggatccga cattgtgatc acacagtcta cagcttcctt aggtgtatct 540  
ctggggcaga gggccaccat ctcatgcagg gccagcaaaa gtgtcagtac atctggctat 600  
agttatatgc actggtacca acagagacca ggacagccac ccaaactcct catctatctt 660  
gcatccaacc tagaatctgg ggtccctgcc aggttcagtg gcagtgggtc tgggacagac 720  
ttcaccctca acatccatcc tgtggaggag gaggatgctg caacctatta ctgtcagcac 780  
agtagggagc ttccgttcac gttcggagggg gggaccaagc tggagataaa acgggctgat 840  
gctgcaccaa ctgtatccac aacaaccctt gccccagac ctctacccc agcccctaca 900  
attgccagcc agcctctgag cctgaggccc gaggcttgta gacctgctgc tggcggagcc 960  
gtgcacacca gaggactgga tttcgctgc gacatctaca tctggcgcc cttggccggg 1020  
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aaactcctgt atatatcaa acaaccattt atgagaccag taaaaactac tcaagaggaa 1140  
gatggctgta gctgccgatt tccagaagaa gaagaaggag gatgtgaact gagagtgaag 1200  
ttcagcagga gcgcagacgc ccccgctac aagcagggcc agaaccagct ctataacgag 1260  
ctcaatctag gacgaagaga ggagtacgat gttttggaca agagacgtgg ccgggaccct 1320

gagatggggg gaaagccgag aaggaagaac cctcaggaag gcctgttcaa tgaactgcag 1380  
aaagataaga tggcggaggc cttcagtgag attgggatga aaggcgagcg ccggaggggc 1440  
aaggggcacg atggcctttt ccagggtctc agtacagcca ccaaggacac cttcgacgcc 1500  
cttcacatgc aggccctgcc ccctcgctga taa 1533

<210> 1619  
<211> 509  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> MNC21XX - muMNC2CAR T841BB3z1XX  
  
<400> 1619

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15  
  
His Ala Ala Arg Pro Glu Val Gln Leu Glu Glu Ser Gly Gly Gly Leu  
20 25 30  
  
Val Lys Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45  
  
Thr Phe Ser Gly Tyr Ala Met Ser Trp Val Arg Gln Thr Pro Glu Lys  
50 55 60  
  
Arg Leu Glu Trp Val Ala Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr  
65 70 75 80  
  
Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95  
  
Lys Asn Thr Leu Tyr Leu Gln Met Ser Ser Leu Arg Ser Glu Asp Thr  
100 105 110  
  
Ala Met Tyr Tyr Cys Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr  
115 120 125  
  
Phe Asp Val Trp Gly Ala Gly Thr Thr Val Thr Val Ser Ser Ala Lys  
130 135 140  
  
Thr Thr Pro Pro Ser Val Tyr Gly Gly Gly Gly Ser Gly Gly Gly Gly  
145 150 155 160  
  
Ser Gly Gly Gly Gly Ser Asp Ile Val Ile Thr Gln Ser Thr Ala Ser  
165 170 175  
  
Leu Gly Val Ser Leu Gly Gln Arg Ala Thr Ile Ser Cys Arg Ala Ser  
180 185 190  
  
Lys Ser Val Ser Thr Ser Gly Tyr Ser Tyr Met His Trp Tyr Gln Gln  
195 200 205

Arg Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr Leu Ala Ser Asn Leu  
 210 215 220

Glu Ser Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp  
 225 230 235 240

Phe Thr Leu Asn Ile His Pro Val Glu Glu Glu Asp Ala Ala Thr Tyr  
 245 250 255

Tyr Cys Gln His Ser Arg Glu Leu Pro Phe Thr Phe Gly Gly Gly Thr  
 260 265 270

Lys Leu Glu Ile Lys Arg Ala Asp Ala Ala Pro Thr Val Ser Thr Thr  
 275 280 285

Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln  
 290 295 300

Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala  
 305 310 315 320

Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala  
 325 330 335

Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr  
 340 345 350

Leu Tyr Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln  
 355 360 365

Pro Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser  
 370 375 380

Cys Arg Phe Pro Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys  
 385 390 395 400

Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln  
 405 410 415

Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu  
 420 425 430

Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg  
 435 440 445

Lys Asn Pro Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met  
 450 455 460

Ala Glu Ala Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly  
 465 470 475 480

Lys Gly His Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp  
485 490 495

Thr Phe Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
500 505

<210> 1620  
<211> 1485  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> huMNC2CAR T841BB3z1XX

<400> 1620  
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gctccaggga aggggctgga gtgggtctca accattagta gtggcggaac ctacatatac 240  
taccccgact cagtgaaggg ccgattcacc atctccagag acaacgcaa gaactcactg 300  
tatctgcaaa tgaacagcct gagagccgag gacacggcgg tgtattactg tgcgagactt 360  
gggggggata attactacga atacttcgat gtctggggca aagggaccac ggtcaccgtc 420  
tcctccggcg gtggcggatc cggcgggtggc ggatccggcg gtggcggatc cgacattgtg 480  
ctgacccagt ctccagcctc cttggccgtg tctccaggac agagggccac catcacctgc 540  
agagccagta agagtgtcag taccagcgga tactcctaca tgcactggta tcagcagaaa 600  
ccaggacaac ctccctaaact cctgattttac ctggcatcca atctggagag cgggggtcca 660  
gccaggttca gcggcagtggt gtctggggacc gatttcaccc tcacaattaa tcctgtggaa 720  
gctaatagata ctgcaaatta ttactgtcag cacagtaggg agctgccttt cacattcggc 780  
ggaggggacca aggtggagat caaacgaact acaacaaccc ctgccccag acctcctacc 840  
ccagccccta caattgccag ccagcctctg agcctgaggc ccgaggcttg tagacctgct 900  
gctggcggag ccgtgcacac cagaggactg gatttcgcct gcgacatcta catctgggcg 960  
cccttggccg ggacttgtgg ggtcctttctc ctgtcactgg ttatcacctt ttactgcaaa 1020  
cggggcagaa agaaactcct gtatatattc aaacaacat ttatgagacc agtacaaaact 1080  
actcaagagg aagatggctg tagctgccga tttccagaag aagaagaagg aggatgtgaa 1140  
ctgagagtga agttcagcag gagcgcagac gccccgcgt acaagcaggg ccagaaccag 1200  
ctctataacg agctcaatct aggacgaaga gaggagtacg atgttttggg caagagacgt 1260  
ggccgggacc ctgagatggg gggaaagccg agaaggaaga accctcagga aggcctgttc 1320  
aatgaactgc agaaagataa gatggcggag gccttcagtg agattgggat gaaaggcgag 1380  
cgccggaggg gcaaggggca cgatggcctt ttccagggtc tcagtacagc caccaaggac 1440  
accttcgacg cccttcacat gcaggccctg cccctcgcgt gataa 1485

<210> 1621  
<211> 493  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> huMNC2CAR T841BB3z1XX

<400> 1621

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Gly Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Gly Leu Glu Trp Val Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr  
65 70 75 80

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95

Lys Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110

Ala Val Tyr Tyr Cys Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr  
115 120 125

Phe Asp Val Trp Gly Lys Gly Thr Thr Val Thr Val Ser Ser Gly Gly  
130 135 140

Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val  
145 150 155 160

Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala  
165 170 175

Thr Ile Thr Cys Arg Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser  
180 185 190

Tyr Met His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu  
195 200 205

Ile Tyr Leu Ala Ser Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser  
210 215 220

Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu  
225 230 235 240

Ala Asn Asp Thr Ala Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro  
245 250 255

Phe Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr  
260 265 270

Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln  
275 280 285

Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala  
290 295 300

Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala  
305 310 315 320

Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr  
325 330 335

Leu Tyr Cys Lys Arg Gly Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln  
340 345 350

Pro Phe Met Arg Pro Val Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser  
355 360 365

Cys Arg Phe Pro Glu Glu Glu Glu Gly Gly Cys Glu Leu Arg Val Lys  
370 375 380

Phe Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln  
385 390 395 400

Leu Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu  
405 410 415

Asp Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg  
420 425 430

Lys Asn Pro Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met  
435 440 445

Ala Glu Ala Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly  
450 455 460

Lys Gly His Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp  
465 470 475 480

Thr Phe Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1622

<211> 1530

<212> DNA

<213> Artificial Sequence



<220>

<223> muMNC2CAR T8283z1XX

<400> 1622

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ctctcctgtg cagcctctgg attcactttc agtggctatg ccatgtcttg ggttcgccag	180
actccggaga agaggctgga gtgggtcgca accattagta gtggtggtac ttatatctac	240
tatccagaca gtgtgaaggg gcgattcacc atctccagag acaatgcaa gaacaccctg	300
tacctgaaaa tgagcagtct gaggtctgag gacacggcca tgtattactg tgcaagactt	360
gggggggata attactacga atacttcgat gtctggggcg cagggaccac ggtcaccgtc	420
tcctccgcca aaacgacacc cccatctgtc tatggcgggt gcggatccgg cggtggcgga	480
tccggcgggt gcggatccga cattgtgatc acacagtcta cagcttcctt aggtgtatct	540
ctggggcaga gggccacat ctcatgcagg gccagcaaaa gtgtcagtac atctggctat	600
agttatatgc actggtacca acagagacca ggacagccac ccaaactcct catctatctt	660
gcatccaacc tagaatctgg ggtccctgcc aggttcagtg gcagtgggtc tgggacagac	720
ttcacctca acatccatcc tgtggaggag gaggatgctg caacctatta ctgtcagcac	780
agtagggagc ttccgttcac gttcggaggg gggaccaagc tggagataaa acgggctgat	840
gctgcaccaa ctgtatccac aacaaccctt gccccagac ctcctacccc agcccctaca	900
attgccagcc agcctctgag cctgaggccc gaggttgta gacctgctgc tggcggagcc	960
gtgcacacca gaggactgga tttcgctgc gacatctaca tctgggcgcc cttggccggg	1020
acttgtaggg tccttctcct gtcactggtt atcacccttt actgcaggag taagaggagc	1080
aggctcctgc acagtgacta catgaacatg actcctagaa gacctgggcc taccagaaag	1140
cattaccagc cctatgcccc accacgcgac ttgcagcct atcgctccag agtgaagttc	1200
agcaggagcg cagacgcccc cgcgtacaag cagggccaga accagctcta taacgagctc	1260
aatctaggac gaagagagga gtacgatgtt ttggacaaga gacgtggccg ggaccctgag	1320
atggggggaa agccgagaag gaagaaccct caggaaggcc tgttcaatga actgcagaaa	1380
gataagatgg cggaggcctt cagtgagatt gggatgaaag gcgagcgccg gaggggcaag	1440
gggcacgatg gccttttcca ggtctcagt acagccacca aggacacctt cgacgccctt	1500
cacatgcagg ccctgcccc tcgctgataa	1530

<210> 1623

<211> 508

<212> PRT

<213> Artificial Sequence

<220>

<223> muMNC2CAR T8283z1XX

<400> 1623

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu

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Pro	Glu	Val	Gln
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Leu	Glu	Glu	Ser
Gly	Gly	Gly	Leu
			30
Val	Lys	Pro	Gly
	35		
Gly	Gly	Ser	Leu
			40
Lys	Leu	Ser	Cys
Ala	Ala	Ser	Gly
			45
Phe			
Thr	Phe	Ser	Gly
	50		
Tyr	Ala	Met	Ser
		55	
Trp	Val	Arg	Gln
			60
Pro	Glu	Lys	
Arg	Leu	Glu	Trp
	65		
Val	Ala	Thr	Ile
	70		
Ser	Ser	Gly	Gly
		75	
Thr	Tyr	Ile	Tyr
			80
Tyr	Pro	Asp	Ser
Val	Lys	Gly	Arg
	85		
Phe	Thr	Ile	Ser
		90	
Arg	Asp	Asn	Ala
		95	
Lys	Asn	Thr	Leu
			100
Tyr	Leu	Gln	Met
			105
Ser	Ser	Leu	Arg
Ser	Glu	Asp	Thr
			110
Ala	Met	Tyr	Tyr
	115		
Cys	Ala	Arg	Leu
			120
Gly	Gly	Asp	Asn
			125
Tyr	Tyr	Glu	Tyr
Phe	Asp	Val	Trp
	130		
Gly	Ala	Gly	Thr
		135	
Thr	Thr	Val	Thr
			140
Val	Ser	Ser	Ala
Lys			
Thr	Thr	Pro	Pro
Ser	Val	Tyr	Gly
	145		
Gly	Gly	Gly	Gly
			150
Ser	Gly	Gly	Gly
			155
Ser	Gly	Gly	Gly
			160
Ser	Gly	Gly	Gly
Gly	Gly	Ser	Asp
			165
Ile	Val	Ile	Thr
			170
Gln	Ser	Thr	Ala
			175
Ser			
Leu	Gly	Val	Ser
			180
Leu	Gly	Gln	Arg
			185
Ala	Thr	Ile	Ser
Cys	Arg	Ala	Ser
			190
Lys	Ser	Val	Ser
Thr	Ser	Gly	Tyr
			195
Ser	Gly	Tyr	Ser
			200
Met	His	Trp	Tyr
			205
Gln	Gln		
Arg	Pro	Gly	Gln
Pro	Pro	Lys	Leu
			210
Leu	Leu	Ile	Tyr
Leu	Ala	Ser	Asn
			215
Leu	Ala	Ser	Asn
			220
Glu	Ser	Gly	Val
Pro	Ala	Arg	Phe
			225
Ser	Gly	Ser	Gly
			230
Ser	Gly	Ser	Gly
			235
Thr	Asp	Ala	Ala
Thr	Tyr		
			240
Phe	Thr	Leu	Asn
Ile	His	Pro	Val
			245
Glu	Glu	Glu	Asp
			250
Ala	Ala	Thr	Tyr
			255
Tyr	Cys	Gln	His
Ser	Arg	Glu	Leu
			260
Pro	Phe	Thr	Phe
Gly	Gly	Gly	Thr
			265
Gly	Gly	Thr	Thr
			270
Lys	Leu	Glu	Ile
Lys	Arg	Ala	Asp
			275
Ala	Ala	Pro	Thr
Val	Ser	Thr	Thr
			280
Val	Ser	Thr	Thr
			285

Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln  
 290 295 300  
 Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala  
 305 310 315 320  
 Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala  
 325 330 335  
 Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Ser Leu Val Ile Thr  
 340 345 350  
 Leu Tyr Cys Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met  
 355 360 365  
 Asn Met Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro  
 370 375 380  
 Tyr Ala Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe  
 385 390 395 400  
 Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu  
 405 410 415  
 Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp  
 420 425 430  
 Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys  
 435 440 445  
 Asn Pro Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala  
 450 455 460  
 Glu Ala Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys  
 465 470 475 480  
 Gly His Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr  
 485 490 495  
 Phe Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
 500 505

<210> 1624  
 <211> 1482  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> huMNC2CAR T8283z1XX

<400> 1624  
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ctctcctgtg cagcctctgg attcaccttc agtggctatg ccatgagctg ggtccgccag	180
gctccaggga aggggctgga gtgggtctca accattagta gtggcggaac ctacatatac	240
taccccgact cagtgaaggg ccgattcacc atctccagag acaacgcaa gaactcactg	300
tatctgcaaa tgaacagcct gagagccgag gacacggccg tgtattactg tgcgagactt	360
gggggggata attactacga atacttcgat gtctggggca aagggaccac ggtcaccgtc	420
tcctccggcg gtggcggtgc cggcggtggc ggatccggcg gtggcggtgc cgacattgtg	480
ctgacccagt ctccagcctc cttggccgtg tctccaggac agagggccac catcacctgc	540
agagccagta agagtgtcag taccagcgga tactcctaca tgcactggta tcagcagaaa	600
ccaggacaac ctctaaact cctgatttac ctggcatcca atctggagag cggggtcca	660
gccaggttca gcggcagtg gtctgggacc gatttcaccc tcacaattaa tcctgtggaa	720
gctaatagata ctgcaaatta ttactgtcag cacagtaggg agctgccttt cacattcggc	780
ggagggacca aggtggagat caaacgaact acaacaacc ctgccccag acctcctacc	840
ccagccccta caattgccag ccagcctctg agcctgaggc ccgaggcttg tagacctgct	900
gctggcggag ccgtgcacac cagaggactg gatttcgcct gcgacatcta catctggcg	960
cccttggccg ggacttgttg ggtccttctc ctgtcactgg ttatcacct ttactgcagg	1020
agtaagagga gcaggctcct gcacagtac tacatgaaca tgactcctag aagacctggg	1080
cctaccagaa agcattacca gccctatgcc ccaccacgc acttcgcagc ctatcgctcc	1140
agagtgaagt tcagcaggag gcgagacgcc cccgcgtaca agcagggcc gaaccagctc	1200
tataacgagc tcaatctagg acgaagagag gactacgatg ttttgacaa gagacgtggc	1260
cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgttcaat	1320
gaactgcaga aagataagat ggcgagggcc ttcagtgaga ttgggatgaa aggcgagcgc	1380
cggaggggca aggggcacga tggccttttc cagggtctca gtacagccac caaggacacc	1440
ttcgacgcc ttcacatgca ggccctgccc cctcgctgat aa	1482

<210> 1625  
 <211> 492  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> huMNC2CAR T8283z1XX

<400> 1625

Met	Ala	Leu	Pro	Val	Thr	Ala	Leu	Leu	Leu	Pro	Leu	Ala	Leu	Leu	Leu
1				5				10					15		

His	Ala	Ala	Arg	Pro	Glu	Val	Gln	Leu	Val	Glu	Ser	Gly	Gly	Gly	Leu
			20				25					30			

Val	Lys	Pro	Gly	Gly	Ser	Leu	Arg	Leu	Ser	Cys	Ala	Ala	Ser	Gly	Phe
		35					40					45			

Thr Phe Ser Gly Tyr Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
 50 55 60

Gly Leu Glu Trp Val Ser Thr Ile Ser Ser Gly Gly Thr Tyr Ile Tyr  
 65 70 75 80

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
 85 90 95

Lys Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
 100 105 110

Ala Val Tyr Tyr Cys Ala Arg Leu Gly Gly Asp Asn Tyr Tyr Glu Tyr  
 115 120 125

Phe Asp Val Trp Gly Lys Gly Thr Thr Val Thr Val Ser Ser Gly Gly  
 130 135 140

Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val  
 145 150 155 160

Leu Thr Gln Ser Pro Ala Ser Leu Ala Val Ser Pro Gly Gln Arg Ala  
 165 170 175

Thr Ile Thr Cys Arg Ala Ser Lys Ser Val Ser Thr Ser Gly Tyr Ser  
 180 185 190

Tyr Met His Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu  
 195 200 205

Ile Tyr Leu Ala Ser Asn Leu Glu Ser Gly Val Pro Ala Arg Phe Ser  
 210 215 220

Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Pro Val Glu  
 225 230 235 240

Ala Asn Asp Thr Ala Asn Tyr Tyr Cys Gln His Ser Arg Glu Leu Pro  
 245 250 255

Phe Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr Thr Thr  
 260 265 270

Thr Pro Ala Pro Arg Pro Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln  
 275 280 285

Pro Leu Ser Leu Arg Pro Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala  
 290 295 300

Val His Thr Arg Gly Leu Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala  
 305 310 315 320

Pro Leu Ala Gly Thr Cys Gly Val Leu Leu Leu Ser Leu Val Ile Thr  
325 330 335

Leu Tyr Cys Arg Ser Lys Arg Ser Arg Leu Leu His Ser Asp Tyr Met  
340 345 350

Asn Met Thr Pro Arg Arg Pro Gly Pro Thr Arg Lys His Tyr Gln Pro  
355 360 365

Tyr Ala Pro Pro Arg Asp Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe  
370 375 380

Ser Arg Ser Ala Asp Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu  
385 390 395 400

Tyr Asn Glu Leu Asn Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp  
405 410 415

Lys Arg Arg Gly Arg Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys  
420 425 430

Asn Pro Gln Glu Gly Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala  
435 440 445

Glu Ala Phe Ser Glu Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys  
450 455 460

Gly His Asp Gly Leu Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr  
465 470 475 480

Phe Asp Ala Leu His Met Gln Ala Leu Pro Pro Arg  
485 490

<210> 1626  
<211> 1467  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> MNE61XX - muMNE6CAR T841BB3z1XX

<400> 1626  
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ctctcctgtg tagtctctgg attcactttc agtagatatg gcatgtcttg ggttcgccag 180  
actccaggca agaggctgga gtgggtcgca accattagtg gtggcggtac ttacatctac 240  
tatccagaca gtgtgaaggg gcgattcacc atctccagag acaatgcaa gaacaccctg 300  
tacctgcaaa tgagcagtct gaagtctgag gacacagcca tgtatcactg tacaagggat 360  
aactacggta ggaactacga ctacggtatg gactactggg gtcaaggaac ctcagtcacc 420  
gtctcctcag gcggtggcgg atccggcggt ggcggtatccg gcggtggcgg atcccaaatt 480

gttctcacc agtctccagc aatcatgtct gcatctccag gggaggaggt caccctaacc 540  
tgcagtgcca cctcaagtgt aagttacata cactggttcc agcagaggcc aggcacttct 600  
cccaaactct ggatttatag cacatccaac ctggcttctg gagtccctgt tcgcttcagt 660  
ggcagtggat atgggacctc ttactctctc acaatcagcc gaatggaggc tgaagatgct 720  
gccacttatt actgccagca aaggagtagt tccccattca cgttcggctc ggggacaaag 780  
ttggaataa aaacaacaac ccctgcccc agacctccta cccagcccc tacaattgcc 840  
agccagcctc tgagcctgag gcccgaggct tgtagacctg ctgctggcgg agccgtgcac 900  
accagaggac tggatttcgc ctgcgacatc tacatctggg cgcccttggc cgggacttgt 960  
ggggtccttc tcctgtcact ggttatcacc ctttactgca aacggggcag aaagaaactc 1020  
ctgtatatat tcaaaacaacc atttatgaga ccagtacaaa ctactcaaga ggaagatggc 1080  
tgtagctgcc gatttccaga agaagaagaa ggaggatgtg aactgagagt gaagttcagc 1140  
aggagcgcag acgccccgc gtacaagcag ggccagaacc agctctataa cgagctcaat 1200  
ctaggacgaa gagaggagta cgatgttttg gacaagagac gtggccggga ccctgagatg 1260  
gggggaaagc cgagaaggaa gaaccctcag gaaggcctgt tcaatgaact gcagaaagat 1320  
aagatggcgg aggccttcag tgagattggg atgaaaggcg agcgccggag gggcaagggg 1380  
cacgatggcc ttttccaggg tctcagtaca gccaccaagg acaccttcga cgcccttcac 1440  
atgcaggccc tgccccctcg ctgataa 1467

<210> 1627  
<211> 487  
<212> PRT  
<213> Artificial Sequence  
  
<220>  
<223> MNE61XX - muMNE6CAR T841BB3z1XX  
  
<400> 1627

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15  
  
His Ala Ala Arg Pro Glu Val Lys Val Val Glu Ser Gly Gly Asp Leu  
20 25 30  
  
Val Lys Pro Gly Gly Ser Leu Lys Leu Ser Cys Val Val Ser Gly Phe  
35 40 45  
  
Thr Phe Ser Arg Tyr Gly Met Ser Trp Val Arg Gln Thr Pro Gly Lys  
50 55 60  
  
Arg Leu Glu Trp Val Ala Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr  
65 70 75 80  
  
Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95

Lys Asn Thr Leu Tyr Leu Gln Met Ser Ser Leu Lys Ser Glu Asp Thr  
100 105 110

Ala Met Tyr His Cys Thr Arg Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr  
115 120 125

Gly Met Asp Tyr Trp Gly Gln Gly Thr Ser Val Thr Val Ser Ser Gly  
130 135 140

Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Ser Gln Ile  
145 150 155 160

Val Leu Thr Gln Ser Pro Ala Ile Met Ser Ala Ser Pro Gly Glu Glu  
165 170 175

Val Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp  
180 185 190

Phe Gln Gln Arg Pro Gly Thr Ser Pro Lys Leu Trp Ile Tyr Ser Thr  
195 200 205

Ser Asn Leu Ala Ser Gly Val Pro Val Arg Phe Ser Gly Ser Gly Tyr  
210 215 220

Gly Thr Ser Tyr Ser Leu Thr Ile Ser Arg Met Glu Ala Glu Asp Ala  
225 230 235 240

Ala Thr Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly  
245 250 255

Ser Gly Thr Lys Leu Glu Ile Lys Thr Thr Thr Pro Ala Pro Arg Pro  
260 265 270

Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser Leu Arg Pro  
275 280 285

Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr Arg Gly Leu  
290 295 300

Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr Cys  
305 310 315 320

Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys Lys Arg Gly  
325 330 335

Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe Met Arg Pro Val  
340 345 350

Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe Pro Glu Glu  
355 360 365



Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser Arg Ser Ala Asp  
370 375 380

Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu Leu Asn  
385 390 395 400

Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg Gly Arg  
405 410 415

Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln Glu Gly  
420 425 430

Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Phe Ser Glu  
435 440 445

Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp Gly Leu  
450 455 460

Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe Asp Ala Leu His  
465 470 475 480

Met Gln Ala Leu Pro Pro Arg  
485

<210> 1628  
<211> 1467  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> huMNE6CAR T841BB3z1XX

<400> 1628  
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ctctcctgtg cagcctctgg attcaccttc agtaggtatg gcatgagctg ggtccgccag 180  
gctccaggga agaggctgga gtgggtctca accattagtg gcggaggcac ctacatatac 240  
taccagact cagtgaaggg ccgattcacc atctccagag acaacgcaa gaacaccctg 300  
tatctgcaaa tgaacagcct gagagccgag gacacggctg tgtattactg taccagagat 360  
aactatggcc gcaactatga ttatggcatg gattattggg gccagggcac cctggtgacc 420  
gtgagcagcg gcggtggcgg atccggcggg ggcggatccg gcggtggcgg atccgaaatt 480  
gtgttgacac agtctccagc caccctgtct ttgtctccag gggaaagagc caccctcacc 540  
tgcagcgcca ccagcagtgt tagctacatc cactggtacc aacagaggcc tggccagagc 600  
cccaggctcc tcatctatag cacctccaac ctggccagcg gcatcccagc caggttcagt 660  
ggcagtgggg ctgggagcga ctacactctc accatcagca gcctagagcc tgaagatttt 720  
gcagtttatt actgtcagca gcgtagcagc tcccccttca cctttggcag cggcaccaaa 780  
gtggaaatta aaacaacaac ccctgcccc agacctccta cccagcccc tacaattgcc 840

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accagaggac tggatttcgc ctgcgacatc tacatctggg cgcccttggc cgggacttgt	960
ggggtccttc tcctgtcact ggttatcacc ctttactgca aacggggcag aaagaaactc	1020
ctgtatatat tcaaaacaacc atttatgaga ccagtacaaa ctactcaaga ggaagatggc	1080
tgtagctgcc gatttccaga agaagaagaa ggaggatgtg aactgagagt gaagttcagc	1140
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ctaggacgaa gagaggagta cgatgttttg gacaagagac gtggccggga ccctgagatg	1260
gggggaaagc cgagaaggaa gaaccctcag gaaggcctgt tcaatgaact gcagaaagat	1320
aagatggcgg aggccttcag tgagattggg atgaaaggcg agcgccggag gggcaagggg	1380
cacgatggcc ttttccaggg tctcagtaca gccaccaagg acaccttcga cgcccttcac	1440
atgcaggccc tgccccctcg ctgataa	1467

<210> 1629

<211> 487

<212> PRT

<213> Artificial Sequence

<220>

<223> huMNE6CAR T841BB3z1XX

<400> 1629

Met	Ala	Leu	Pro	Val	Thr	Ala	Leu	Leu	Leu	Pro	Leu	Ala	Leu	Leu	Leu
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His	Ala	Ala	Arg	Pro	Glu	Val	Gln	Leu	Val	Glu	Ser	Gly	Gly	Gly	Leu
			20					25					30		

Val	Lys	Pro	Gly	Gly	Ser	Leu	Arg	Leu	Ser	Cys	Ala	Ala	Ser	Gly	Phe
		35					40					45			

Thr	Phe	Ser	Arg	Tyr	Gly	Met	Ser	Trp	Val	Arg	Gln	Ala	Pro	Gly	Lys
	50					55					60				

Arg	Leu	Glu	Trp	Val	Ser	Thr	Ile	Ser	Gly	Gly	Gly	Thr	Tyr	Ile	Tyr
65					70					75				80	

Tyr	Pro	Asp	Ser	Val	Lys	Gly	Arg	Phe	Thr	Ile	Ser	Arg	Asp	Asn	Ala
				85					90					95	

Lys	Asn	Thr	Leu	Tyr	Leu	Gln	Met	Asn	Ser	Leu	Arg	Ala	Glu	Asp	Thr
		100						105					110		

Ala	Val	Tyr	Tyr	Cys	Thr	Arg	Asp	Asn	Tyr	Gly	Arg	Asn	Tyr	Asp	Tyr
		115					120					125			

Gly	Met	Asp	Tyr	Trp	Gly	Gln	Gly	Thr	Leu	Val	Thr	Val	Ser	Ser	Gly
130						135					140				

Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Ile  
145 150 155 160

Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly Glu Arg  
165 170 175

Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp  
180 185 190

Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr Ser Thr  
195 200 205

Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser  
210 215 220

Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe  
225 230 235 240

Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly  
245 250 255

Ser Gly Thr Lys Val Glu Ile Lys Thr Thr Thr Pro Ala Pro Arg Pro  
260 265 270

Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser Leu Arg Pro  
275 280 285

Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr Arg Gly Leu  
290 295 300

Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr Cys  
305 310 315 320

Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys Lys Arg Gly  
325 330 335

Arg Lys Lys Leu Leu Tyr Ile Phe Lys Gln Pro Phe Met Arg Pro Val  
340 345 350

Gln Thr Thr Gln Glu Glu Asp Gly Cys Ser Cys Arg Phe Pro Glu Glu  
355 360 365

Glu Glu Gly Gly Cys Glu Leu Arg Val Lys Phe Ser Arg Ser Ala Asp  
370 375 380

Ala Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu Leu Asn  
385 390 395 400

Leu Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg Gly Arg  
405 410 415

Asp Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln Glu Gly

Leu Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Phe Ser Glu  
 435 440 445

Ile Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp Gly Leu  
 450 455 460

Phe Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe Asp Ala Leu His  
 465 470 475 480

Met Gln Ala Leu Pro Pro Arg  
 485

<210> 1630

<211> 1464

<212> DNA

<213> Artificial Sequence

<220>

<223> muMNE6CAR T84283z1XX

<400> 1630

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ctctcctgtg tagtctctgg attcactttc agtagatatg gcatgtcttg ggttcgccag	180
actccaggca agaggctgga gtgggtcgca accattagtg gtggcggtag ttacatctac	240
tatccagaca gtgtgaaggg gcgattcacc atctccagag acaatgcaa gaacaccctg	300
tacctgaaa tgagcagtct gaagtctgag gacacagcca tgtatcactg tacaagggat	360
aactacggtg ggaactacga ctacggtatg gactactggg gtcaaggaa ctcagtcacc	420
gtctcctcag gcggtggcgg atccggcggg ggcggatccg gcggtggcgg atcccaaatt	480
gttctcacc agtctccagc aatcatgtct gcatctccag gggaggaggt caccctaacc	540
tgcagtgcc cctcaagtgt aagttacata cactggttcc agcagaggcc aggcacttct	600
cccaaactct ggatttatag cacatccaac ctggcttctg gaggccctgt tcgcttcagt	660
ggcagtggat atgggacctt ttactctctc acaatcagcc gaatggaggc tgaagatgct	720
gccacttatt actgccagca aaggagtagt tccccattca cgttcggctc ggggacaaag	780
ttggaaataa aaacaacaac cctgcccc agacctccta cccagcccc tacaattgcc	840
agccagcctc tgagcctgag gcccagggtg tgtagacctg ctgctggcgg agccgtgcac	900
accagaggac tggatttcgc ctgcgacatc tacatctggg cgcccttggc cgggacttgt	960
ggggtccttc tcctgtcact ggttatcacc ctttactgca ggagtaagag gagcaggctc	1020
ctgcacagtg actacatgaa catgactcct agaagacctg ggcctaccag aaagcattac	1080
cagccctatg cccaccacg cgacttcgca gcctatcgct ccagagtga gttcagcagg	1140
agcgagacg ccccgcgta caagcagggc cagaaccagc tctataacga gctcaatcta	1200
ggacgaagag aggagtacga tgttttggac aagagacgtg gccgggaccc tgagatgggg	1260

ggaaagccga gaaggaagaa ccctcaggaa ggcctgttca atgaactgca gaaagataag	1320
atggcggagg ccttcagtga gattgggatg aaaggcgagc gccggagggg caaggggcac	1380
gatggccttt tccaggtct cagtacagcc accaaggaca ctttcgacgc cttcacatg	1440
caggccctgc cccctcgctg ataa	1464

<210> 1631  
 <211> 486  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> muMNE6CAR T84283z1XX

<400> 1631

Met	Ala	Leu	Pro	Val	Thr	Ala	Leu	Leu	Leu	Pro	Leu	Ala	Leu	Leu	Leu
1				5				10					15		

His	Ala	Ala	Arg	Pro	Glu	Val	Lys	Val	Val	Glu	Ser	Gly	Gly	Asp	Leu
			20				25						30		

Val	Lys	Pro	Gly	Gly	Ser	Leu	Lys	Leu	Ser	Cys	Val	Val	Ser	Gly	Phe
		35					40					45			

Thr	Phe	Ser	Arg	Tyr	Gly	Met	Ser	Trp	Val	Arg	Gln	Thr	Pro	Gly	Lys
	50					55					60				

Arg	Leu	Glu	Trp	Val	Ala	Thr	Ile	Ser	Gly	Gly	Gly	Thr	Tyr	Ile	Tyr
65					70					75					80

Tyr	Pro	Asp	Ser	Val	Lys	Gly	Arg	Phe	Thr	Ile	Ser	Arg	Asp	Asn	Ala
			85						90					95	

Lys	Asn	Thr	Leu	Tyr	Leu	Gln	Met	Ser	Ser	Leu	Lys	Ser	Glu	Asp	Thr
			100					105					110		

Ala	Met	Tyr	His	Cys	Thr	Arg	Asp	Asn	Tyr	Gly	Arg	Asn	Tyr	Asp	Tyr
		115					120					125			

Gly	Met	Asp	Tyr	Trp	Gly	Gln	Gly	Thr	Ser	Val	Thr	Val	Ser	Ser	Gly
	130					135					140				

Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser	Gln	Ile
145					150					155					160

Val	Leu	Thr	Gln	Ser	Pro	Ala	Ile	Met	Ser	Ala	Ser	Pro	Gly	Glu	Glu
				165					170					175	

Val	Thr	Leu	Thr	Cys	Ser	Ala	Thr	Ser	Ser	Val	Ser	Tyr	Ile	His	Trp
			180					185					190		

Phe	Gln	Gln	Arg	Pro	Gly	Thr	Ser	Pro	Lys	Leu	Trp	Ile	Tyr	Ser	Thr
-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----

195

200

205

Ser	Asn	Leu	Ala	Ser	Gly	Val	Pro	Val	Arg	Phe	Ser	Gly	Ser	Gly	Tyr
210						215					220				
Gly	Thr	Ser	Tyr	Ser	Leu	Thr	Ile	Ser	Arg	Met	Glu	Ala	Glu	Asp	Ala
225					230					235					240
Ala	Thr	Tyr	Tyr	Cys	Gln	Gln	Arg	Ser	Ser	Ser	Pro	Phe	Thr	Phe	Gly
				245					250					255	
Ser	Gly	Thr	Lys	Leu	Glu	Ile	Lys	Thr	Thr	Thr	Pro	Ala	Pro	Arg	Pro
			260					265					270		
Pro	Thr	Pro	Ala	Pro	Thr	Ile	Ala	Ser	Gln	Pro	Leu	Ser	Leu	Arg	Pro
		275					280					285			
Glu	Ala	Cys	Arg	Pro	Ala	Ala	Gly	Gly	Ala	Val	His	Thr	Arg	Gly	Leu
	290					295					300				
Asp	Phe	Ala	Cys	Asp	Ile	Tyr	Ile	Trp	Ala	Pro	Leu	Ala	Gly	Thr	Cys
305					310					315					320
Gly	Val	Leu	Leu	Leu	Ser	Leu	Val	Ile	Thr	Leu	Tyr	Cys	Arg	Ser	Lys
				325					330						335
Arg	Ser	Arg	Leu	Leu	His	Ser	Asp	Tyr	Met	Asn	Met	Thr	Pro	Arg	Arg
			340					345					350		
Pro	Gly	Pro	Thr	Arg	Lys	His	Tyr	Gln	Pro	Tyr	Ala	Pro	Pro	Arg	Asp
		355					360					365			
Phe	Ala	Ala	Tyr	Arg	Ser	Arg	Val	Lys	Phe	Ser	Arg	Ser	Ala	Asp	Ala
	370					375					380				
Pro	Ala	Tyr	Lys	Gln	Gly	Gln	Asn	Gln	Leu	Tyr	Asn	Glu	Leu	Asn	Leu
385					390					395					400
Gly	Arg	Arg	Glu	Glu	Tyr	Asp	Val	Leu	Asp	Lys	Arg	Arg	Gly	Arg	Asp
				405					410					415	
Pro	Glu	Met	Gly	Gly	Lys	Pro	Arg	Arg	Lys	Asn	Pro	Gln	Glu	Gly	Leu
			420					425					430		
Phe	Asn	Glu	Leu	Gln	Lys	Asp	Lys	Met	Ala	Glu	Ala	Phe	Ser	Glu	Ile
	435						440					445			
Gly	Met	Lys	Gly	Glu	Arg	Arg	Arg	Gly	Lys	Gly	His	Asp	Gly	Leu	Phe
	450					455					460				
Gln	Gly	Leu	Ser	Thr	Ala	Thr	Lys	Asp	Thr	Phe	Asp	Ala	Leu	His	Met
465					470					475					480

Gln Ala Leu Pro Pro Arg  
485

<210> 1632  
<211> 1464  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> huMNE6CAR T84283z1XX

<400> 1632  
atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg 60  
ccggagggtgc agctggtgga gtctggggga ggcctggtca agcctggggg gtccctgaga 120  
ctctcctgtg cagcctctgg attcaccttc agtaggtatg gcatgagctg ggtccgccag 180  
gctccaggga agaggctgga gtgggtctca accattagt ggcgaggcac ctacatatac 240  
taccagact cagtgaaggg ccgattcacc atctccagag acaacgcaa gaacaccctg 300  
tatctgcaa tgaacagcct gagagccgag gacacggctg tgtattactg taccagagat 360  
aactatggcc gcaactatga ttatggcatg gattattggg gccagggcac cctggtgacc 420  
gtgagcagcg gcggtggcgg atccggcggg ggcggatccg gcggtggcgg atccgaaatt 480  
gtgttgacac agtctccagc caccctgtct ttgtctccag gggaaagagc caccctcacc 540  
tgcagcgcca ccagcagtgt tagctacatc cactggtacc aacagaggcc tggccagagc 600  
cccaggctcc tcatttatag cacctccaac ctggccagcg gcatcccagc caggttcagt 660  
ggcagtgggt ctgggagcga ctacactctc accatcagca gcctagagcc tgaagatttt 720  
gcagttttatt actgtcagca gcgtagcagc tcccccttca cctttggcag cggcaccaaa 780  
gtggaaatta aaacaacaac ccctgcccc agacctccta cccagcccc tacaattgcc 840  
agccagcctc tgagcctgag gcccagggtc ttagacctg ctgctggcgg agccgtgcac 900  
accagaggac tggatttcgc ctgcgacatc tacatctggg cgcccttggc cgggacttgt 960  
ggggtccttc tcctgtcact ggttatcacc ctttactgca ggagtaagag gagcaggctc 1020  
ctgcacagtg actacatgaa catgactcct agaagacctg ggcctaccag aaagcattac 1080  
cagccctatg cccaccacg cgacttcgca gcctatcgct ccagagtga gttcagcagg 1140  
agcgagacg ccccgcgta caagcagggc cagaaccagc tctataacga gctcaatcta 1200  
ggacgaagag aggagtacga tgttttgac aagagacgtg gccgggaccc tgagatgggg 1260  
ggaaagccga gaaggaagaa cctcaggaa ggcctgttca atgaactgca gaaagataag 1320  
atggcgaggg cttcagtga gattgggatg aaaggcgagc gccggagggg caaggggcac 1380  
gatggccttt tccagggtct cagtacagcc accaaggaca ctttcgacgc cttcacatg 1440  
caggccctgc cccctcgctg ataa 1464

<210> 1633  
<211> 486  
<212> PRT

<213> Artificial Sequence

<220>

<223> huMNE6CAR T84283z1XX

<400> 1633

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu  
20 25 30

Val Lys Pro Gly Gly Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe  
35 40 45

Thr Phe Ser Arg Tyr Gly Met Ser Trp Val Arg Gln Ala Pro Gly Lys  
50 55 60

Arg Leu Glu Trp Val Ser Thr Ile Ser Gly Gly Gly Thr Tyr Ile Tyr  
65 70 75 80

Tyr Pro Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala  
85 90 95

Lys Asn Thr Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr  
100 105 110

Ala Val Tyr Tyr Cys Thr Arg Asp Asn Tyr Gly Arg Asn Tyr Asp Tyr  
115 120 125

Gly Met Asp Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly  
130 135 140

Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Ser Glu Ile  
145 150 155 160

Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly Glu Arg  
165 170 175

Ala Thr Leu Thr Cys Ser Ala Thr Ser Ser Val Ser Tyr Ile His Trp  
180 185 190

Tyr Gln Gln Arg Pro Gly Gln Ser Pro Arg Leu Leu Ile Tyr Ser Thr  
195 200 205

Ser Asn Leu Ala Ser Gly Ile Pro Ala Arg Phe Ser Gly Ser Gly Ser  
210 215 220

Gly Ser Asp Tyr Thr Leu Thr Ile Ser Ser Leu Glu Pro Glu Asp Phe  
225 230 235 240

Ala Val Tyr Tyr Cys Gln Gln Arg Ser Ser Ser Pro Phe Thr Phe Gly  
245 250 255



Ser Gly Thr Lys Val Glu Ile Lys Thr Thr Thr Pro Ala Pro Arg Pro  
 260 265 270

Pro Thr Pro Ala Pro Thr Ile Ala Ser Gln Pro Leu Ser Leu Arg Pro  
 275 280 285

Glu Ala Cys Arg Pro Ala Ala Gly Gly Ala Val His Thr Arg Gly Leu  
 290 295 300

Asp Phe Ala Cys Asp Ile Tyr Ile Trp Ala Pro Leu Ala Gly Thr Cys  
 305 310 315 320

Gly Val Leu Leu Leu Ser Leu Val Ile Thr Leu Tyr Cys Arg Ser Lys  
 325 330 335

Arg Ser Arg Leu Leu His Ser Asp Tyr Met Asn Met Thr Pro Arg Arg  
 340 345 350

Pro Gly Pro Thr Arg Lys His Tyr Gln Pro Tyr Ala Pro Pro Arg Asp  
 355 360 365

Phe Ala Ala Tyr Arg Ser Arg Val Lys Phe Ser Arg Ser Ala Asp Ala  
 370 375 380

Pro Ala Tyr Lys Gln Gly Gln Asn Gln Leu Tyr Asn Glu Leu Asn Leu  
 385 390 395 400

Gly Arg Arg Glu Glu Tyr Asp Val Leu Asp Lys Arg Arg Gly Arg Asp  
 405 410 415

Pro Glu Met Gly Gly Lys Pro Arg Arg Lys Asn Pro Gln Glu Gly Leu  
 420 425 430

Phe Asn Glu Leu Gln Lys Asp Lys Met Ala Glu Ala Phe Ser Glu Ile  
 435 440 445

Gly Met Lys Gly Glu Arg Arg Arg Gly Lys Gly His Asp Gly Leu Phe  
 450 455 460

Gln Gly Leu Ser Thr Ala Thr Lys Asp Thr Phe Asp Ala Leu His Met  
 465 470 475 480

Gln Ala Leu Pro Pro Arg  
 485

<210> 1634

<211> 117

<212> DNA

<213> Artificial Sequence

<220>

<223> Minimal CMV promoter (mCMV)

<400> 1634  
taggcgtgta cgggtgggagg cctatataag cagagctcgt ttagtgaacc gtcagatcgc 60  
ctggagacgc catccacgct gttttgacct ccatagaaga caccgggacc gatccag 117

<210> 1635  
<211> 114  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Minimal IL2 promoter (mIL2P)

<400> 1635  
cattttgaca cccccataat atttttccag aattaacagt ataaattgca tctcttggtc 60  
aagagttccc tatcactctc tttaatcact actcacagta acctcaactc ctgc 114

<210> 1636  
<211> 31  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Minimal promoter, miniP

<400> 1636  
agagggtata taatggaagc tcgacttcca g 31

<210> 1637  
<211> 540  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> IL18 activated

<400> 1637  
atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg 60  
ccgtacttcg gcaagctgga aagcaagctg agcgtgatcc ggaacctgaa cgaccaggtg 120  
ctgttcacgc atcagggcaa cagaccctg ttcgaggaca tgaccgacag cgactgcaga 180  
gacaacgccc ctcggacat cttcatcatc agcatgtaca aggacagcca gcctagaggc 240  
atggccgtga ccatctctgt gaagtgcgag aagatcagca ccctgagctg cgagaacaag 300  
atcatcagct tcaaagagat gaacccgccg gacaacatca aggacaccaa gagcgacatc 360  
atatctttcc agcggagcgt gccgggccac gacaacaaga tgcagtttga gagcagcagc 420  
tacgagggct acttcctggc ctgcgagaaa gagcgggacc tgttcaagct gacacctgaag 480  
aaagaggacg aactgggcga ccgcagcatc atgttcaccg tgcagaacga ggactgataa 540

<210> 1638  
<211> 178  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> IL18 activated

<400> 1638

Met Ala Leu Pro Val Thr Ala Leu Leu Leu Pro Leu Ala Leu Leu Leu  
1 5 10 15

His Ala Ala Arg Pro Tyr Phe Gly Lys Leu Glu Ser Lys Leu Ser Val  
20 25 30

Ile Arg Asn Leu Asn Asp Gln Val Leu Phe Ile Asp Gln Gly Asn Arg  
35 40 45

Pro Leu Phe Glu Asp Met Thr Asp Ser Asp Cys Arg Asp Asn Ala Pro  
50 55 60

Arg Thr Ile Phe Ile Ile Ser Met Tyr Lys Asp Ser Gln Pro Arg Gly  
65 70 75 80

Met Ala Val Thr Ile Ser Val Lys Cys Glu Lys Ile Ser Thr Leu Ser  
85 90 95

Cys Glu Asn Lys Ile Ile Ser Phe Lys Glu Met Asn Pro Pro Asp Asn  
100 105 110

Ile Lys Asp Thr Lys Ser Asp Ile Ile Phe Phe Gln Arg Ser Val Pro  
115 120 125

Gly His Asp Asn Lys Met Gln Phe Glu Ser Ser Ser Tyr Glu Gly Tyr  
130 135 140

Phe Leu Ala Cys Glu Lys Glu Arg Asp Leu Phe Lys Leu Ile Leu Lys  
145 150 155 160

Lys Glu Asp Glu Leu Gly Asp Arg Ser Ile Met Phe Thr Val Gln Asn  
165 170 175

Glu Asp

<210> 1639

<211> 770

<212> DNA

<213> Artificial Sequence

<220>

<223> Foxp3NFATc1X3IL18

<400> 1639

ggcttcattt tttccattta ctgcagaggc ttcatttttt ccattttactg cagaggcttc 60

atTTTTtcca tttactgcag aactagtttag gcgtgtacgg tgggaggcct atataagcag 120

agctcgttta gtgaaccgtc agatcgcttg gagacgcat ccacgctgtt ttgacctcca 180

tagaagacac cgggaccgat ccagcctcga gagaccaat gctagccacc atggccttac 240

cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg ccgtacttcg 300

gcaagctgga aagcaagctg agcgtgatcc ggaacctgaa cgaccaggtg ctgttcatcg	360
atcagggcaa cagacccttg ttcgaggaca tgaccgacag cgactgcaga gacaacgccc	420
ctcggaccat cttcatcatc agcatgtaca aggacagcca gcctagaggc atggccgtga	480
ccatctctgt gaagtgcgag aagatcagca ccctgagctg cgagaacaag atcatcagct	540
tcaaagagat gaacccgccg gacaacatca aggacaccaa gagcgacatc atattcttcc	600
agcggagcgt gcccggccac gacaacaaga tgcagtttga gagcagcagc tacgagggt	660
acttcctggc ctgcgagaaa gagcgggacc tgttcaagct gatcctgaag aaagaggacg	720
aactgggcga ccgcagcatc atgttcaccg tgcagaacga ggactgataa	770

<210> 1640  
 <211> 850  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Foxp3NFATc1X6IL18

<400> 1640	
gcttcatttt ttccatttac tgcagaggct tcattttttc cattttactgc agaggcttca	60
ttttttccat ttactgcaga ggcttcattt tttccattta ctgcagaggc ttcatTTTTT	120
ccattttactg cagaggcttc atttttttcca tttactgcag aactagttag gcgtgtacgg	180
tgggaggcct atataagcag agctcgttta gtgaaccgtc agatcgcctg gagacgcat	240
ccacgctggt ttgacctcca tagaagacac cgggaccgat ccagcctcga gagaccaat	300
gctagccacc atggccttac cagtgaccgc ctgtctcctg ccgctggcct tgctgctcca	360
cgccgccagg ccgtacttcg gcaagctgga aagcaagctg agcgtgatcc ggaacctgaa	420
cgaccaggtg ctgttcatcg atcagggcaa cagacccttg ttcgaggaca tgaccgacag	480
cgactgcaga gacaacgccc ctcggaccat cttcatcatc agcatgtaca aggacagcca	540
gcctagaggc atggccgtga ccatctctgt gaagtgcgag aagatcagca ccctgagctg	600
cgagaacaag atcatcagct tcaaagagat gaacccgccg gacaacatca aggacaccaa	660
gagcgacatc atattcttcc agcggagcgt gcccggccac gacaacaaga tgcagtttga	720
gagcagcagc tacgagggt acttcctggc ctgcgagaaa gagcgggacc tgttcaagct	780
gatcctgaag aaagaggacg aactgggcga ccgcagcatc atgttcaccg tgcagaacga	840
ggactgataa	850

<210> 1641  
 <211> 779  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> IL2NFATc1X3IL18

<400> 1641	
ggaggaaaaa ctgtttcata cagaaggcgt ggaggaaaaa ctgtttcata cagaaggcgt	60
ggaggaaaaa ctgtttcata cagaaggcgt actagttagg cgtgtacggt gggaggccta	120

tataagcaga gctcgttttag tgaaccgtca gatcgcttg agacgccatc cacgctgttt	180
tgacctccat agaagacacc gggaccgatc cagcctcgag agaccaatg ctagccacca	240
tggccttacc agtgaccgcc ttgctcctgc cgctggcctt gctgctccac gccgccaggc	300
cgtacttcgg caagctggaa agcaagctga gcgtgatccg gaacctgaac gaccaggtgc	360
tgttcatcga tcagggaac agaccctgt tcgaggacat gaccgacagc gactgcagag	420
acaacgcccc tcggaccatc ttcacatca gcatgtacaa ggacagccag cctagaggca	480
tggccgtgac catctctgtg aagtgcgaga agatcagcac cctgagctgc gagaacaaga	540
tcacagctt caaagagatg aacccgccgg acaacatcaa ggacaccaag agcgacatca	600
tattcttcca gcggagcgtg cccggccacg acaacaagat gcagtttgag agcagcagct	660
acgagggcta cttcctggcc tgcgagaaag agcgggacct gttcaagctg atcctgaaga	720
aagaggacga actgggcgac cgcagcatca tgttcaccgt gcagaacgag gactgataa	779

<210> 1642  
 <211> 868  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> IL2NFATc1X6IL18

<400> 1642	
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ggaggaaaaa ctgtttcata cagaaggcgt ggaggaaaaa ctgtttcata cagaaggcgt	120
ggaggaaaaa ctgtttcata cagaaggcgt ggaggaaaaa ctgtttcata cagaaggcgt	180
actagttagg cgtgtacggt gggaggccta tataagcaga gctcgttttag tgaaccgtca	240
gatcgcttg agacgccatc cacgctgttt tgacctccat agaagacacc gggaccgatc	300
cagctcgaga gaccaatgc tagccacat ggccctacca gtgaccgcct tgctcctgcc	360
gctggccttg ctgctccacg ccgccaggcc gtacttcggc aagctggaaa gcaagctgag	420
cgtgatccgg aacctgaacg accaggtgct gttcatcgat cagggaaca gacctgtt	480
cgaggacatg accgacagc actgcagaga caacgcccct cggaccatct tcacatcag	540
catgtacaag gacagccagc ctagaggcat ggccgtgacc atctctgtga agtgcgagaa	600
gatcagcacc ctgagctgcg agaacaagat catcagcttc aaagagatga acccgccgga	660
caacatcaag gacaccaaga gcgacatcat attcttcag cggagcgtgc ccggccacga	720
caacaagatg cagtttgaga gcagcagcta cgagggtac ttcttgccct gcgagaaaga	780
gcgggacctg ttcaagctga tcctgaagaa agaggacgaa ctgggcgacc gcagcatcat	840
gttcaccgtg cagaacgagg actgataa	868

<210> 1643  
 <211> 2347  
 <212> DNA  
 <213> Artificial Sequence

<220>

<223> Best antibodies with inducible IL18 +/- 1XX mutations in ITAMs of  
CD3zeta - MNC2 - muMNC2841BB3zFoxp3NFATIL18

<400> 1643

atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg	60
ccggagggtcc agctggagga gtcaggggga ggcttagtga agcctggagg gtccctgaaa	120
ctctcctgtg cagcctctgg attcactttc agtggctatg ccatgtcttg ggttcgccag	180
actccggaga agaggctgga gtgggtcgca accattagta gtggtggtac ttatatctac	240
tatccagaca gtgtgaaggg gcgattcacc atctccagag acaatgcaa gaacaccctg	300
tacctgaaa tgagcagtct gaggtctgag gacacggcca tgtattactg tgcaagactt	360
gggggggata attactacga atacttcgat gtctggggcg cagggaccac ggtcaccgtc	420
tcctccgcca aaacgacacc cccatctgtc tatggcgggt gcggatccgg cggtggcgga	480
tccggcgggt gcggatccga cattgtgatc acacagtcta cagcttcctt aggtgtatct	540
ctggggcaga gggccacat ctcatgcagg gccagcaaaa gtgtcagtac atctggctat	600
agttatatgc actggtacca acagagacca ggacagccac ccaaactcct catctatctt	660
gcatccaacc tagaatctgg ggtccctgcc aggttcagtg gcagtgggtc tgggacagac	720
ttcacctca acatccatcc tgtggaggag gaggatgctg caacctatta ctgtcagcac	780
agtagggagc ttccgttcac gttcggaggg gggaccaagc tggagataaa acgggctgat	840
gctgcaccaa ctgtatccac aacaaccctt gccccagac ctctacccc agcccctaca	900
attgccagcc agcctctgag cctgaggccc gaggttgta gacctgctgc tggcggagcc	960
gtgcacacca gaggactgga tttcgctgc gacatctaca tctgggcgcc cttggccggg	1020
acttgtaggg tccttctcct gtcactggtt atcacctttt actgcaaagc gggcagaaag	1080
aaactcctgt atatatcaa acaaccattt atgagaccag taaaaactac tcaagaggaa	1140
gatggctgta gctgccgatt tccagaagaa gaagaaggag gatgtgaact gagagtgaag	1200
ttcagcagga gcgcagacgc ccccgctac aagcagggcc agaaccagct ctataacgag	1260
ctcaatctag gacgaagaga ggagtacgat gttttggaca agagacgtgg ccgggaccct	1320
gagatggggg gaaagccgag aaggaagaac cctcaggaag gcctgtacaa tgaactgcag	1380
aaagataaga tggcggaggc ctacagttag attgggatga aaggcgagcg ccggaggggc	1440
aaggggcacg atggccttta ccagggtctc agtacagcca ccaaggacac ctacgacgcc	1500
cttcacatgc aggccctgcc cctcgtctga taagtttaaa ctgccagaac atttctctgg	1560
cctaactggc cgttaccggc ttcatTTTTT ccatttactg cagaggcttc atTTTTTcca	1620
tttactgcag aggccttatt ttttccattt actgcagaac tagttaggcg tgtacggtgg	1680
gaggcctata taagcagagc tcgttttagtg aaccgtcaga tcgcctggag acgccatcca	1740
cgctgttttg acctccatag aagacaccgg gaccgatcca gcctcgagag acccaatgct	1800
agccaccatg gccttaccag tgaccgcctt gctcctgccg ctggccttgc tgctccacgc	1860
cgccaggccg tacttcggca agctggaaa gcaagctgagc gtgatccgga acctgaacga	1920

ccaggtgctg ttcatcgatc agggcaacag acccctgttc gaggacatga cgcacagcga	1980
ctgcagagac aacgcccctc ggaccatctt catcatcagc atgtacaagg acagccagcc	2040
tagaggcatg gccgtgacca tctctgtgaa gtgcgagaag atcagcacc ttagctgcga	2100
gaacaagatc atcagcttca aagagatgaa cccgccggac aacatcaagg acaccaagag	2160
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<212> DNA

<213> Artificial Sequence

<220>

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<211> 2344

<212> DNA

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cgagggttac ttcctggcct gcgagaaaga gcgggacctg ttcaagctga tcctgaagaa	2220
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<211> 2281

<212> DNA

<213> Artificial Sequence

<220>

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<400> 1655

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cccaaactct ggatttatag cacatccaac ctggcttctg gagtccctgt tcgcttcagt	660
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ctacgagggc tacttcctgg cctgcgagaa agagcgggac ctgttcaagc tgatcctgaa	2220
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 <211> 2281  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> huMNE6841BB3z1XXFoxp3NFATIL18

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gcagtttatt actgtcagca gcgtagcagc tccccctttca cctttggcag cggcaccaaa	780
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ctacgagggc tacttcctgg cctgcgagaa agagcgggac ctgttcaagc tgatcctgaa	2220
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<210> 1657  
 <211> 2278  
 <212> DNA  
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<220>  
 <223> muMNE68283z1XXFoxp3NFATIL18

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actccaggca agaggctgga gtgggtcgca accattagtg gtggcggtag ttacatctac	240
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gccacttatt actgccagca aaggagtagt tccccattca cgttcggctc ggggacaaag	780
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<210> 1658  
 <211> 2278  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> huMNE68283z1XXFoxp3NFATIL18

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gctccaggga agaggctgga gtgggtctca accattagt gcggaggcac ctacatatac	240
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<210> 1659  
 <211> 2293  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> mu20A10841BB3zFoxp3NFATIL18

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<210> 1660  
 <211> 2296  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> hu20A10841BB3zFxp3NFATIL18

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<211> 2290

<212> DNA

<213> Artificial Sequence

<220>

<223> mu20A108283zFoxp3NFATIL18

<400> 1661

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 <212> DNA  
 <213> Artificial Sequence

<220>  
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<211> 2293

<212> DNA

<213> Artificial Sequence

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<400> 1663

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<211> 2296

<212> DNA

<213> Artificial Sequence

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<400> 1664

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<212> DNA

<213> Artificial Sequence

<220>

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<400> 1666

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<400> 1667

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<400> 1668

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cgactgcaga gacaacgccc ctcggaccat cttcatcatc agcatgtaca aggacagcca	1980
gcctagaggc atggccgtga ccattcttgt gaagtgcgag aagatcagca ccctgagctg	2040
cgagaacaag atcatcagct tcaaagagat gaacccgccg gacaacatca aggacacaa	2100
gagcgacatc atattcttcc agcggagcgt gcccggccac gacaacaaga tgcagtttga	2160
gagcagcagc tacgagggtt acttcctggc ctgcgagaaa gagcgggacc tgttcaagct	2220
gatcctgaag aaagaggacg aactgggcga ccgcagcatc atgttcaccg tgcagaacga	2280
ggactgataa	2290

<210> 1672  
 <211> 2290  
 <212> DNA  
 <213> Artificial Sequence

<220>

<223> hu25E6841BB3z1XXFoxp3NFATIL18

<400> 1672

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ctgagctgcg ccgccagcgg cttcacattc agcagctacg gcatgagctg ggtgcggcag	180
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taccccgaca gcgtgaaggg cagattcacc atctcaagag ataacgcaa gaacagcctg	300
tacctgcaga tgaacagcct gcggggccgag gacaccgccg tgtactactg cgccagacag	360
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<210> 1673  
 <211> 2287  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> mu25E68283z1XXFxp3NFATIL18

<400> 1673	
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ctctcctgtg cagcctctgg tttcactttc agtagttatg gaatgtcttg ggttcgccag	180
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cagactccac tcactttgtc ggttaccatt ggacaaccag cctccatctc ttgcaagtca	540
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acaattgcc a gccagcctct gagcctgagg cccgaggctt gtagacctgc tgctggcgga	900
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ctgataa	2287

<210> 1674  
 <211> 2287  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> hu25E68283z1XXFoxp3NFATIL18

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ctgagctgcg ccgccagcgg cttcacattc agcagctacg gcatgagctg ggtgcggcag	180
gcccctggca agggcctgga atgggtcagc accatcagca acggcggaag acacaccttc	240
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ctgataa						2287

<210> 1675  
 <211> 1482  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> pCDH MSCV h20A10-0 CAR 41BB

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ctgagctgtg	ccgccagcgg	cttcaccttt agcacatacg ccatgagctg ggtccgacag 180
gccccctggaa	aaggccttga	atgggttgcc tctatcgga gagccggcag cacctactac 240

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aagcccggcc agcctcctaa gctgctgata tactggggcca gcaccagaga aagcggcgtg	660
cccgatagat tttctggcag cggctctggc accgacttca ccctgacaat tagctccctg	720
caggccgagg atgtggccgt gtactactgt caccagtacc tgagcagcct gacctttggc	780
ggcggaaaca aggtggaaat caagcgaaca acaaccctg ccccagacc tcctaccca	840
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gaactgcaga aagataagat ggccgaggcc tacagtgaga ttgggatgaa aggcgagcgc	1380
cggaggggca aggggcacga tggcctttac cagggtctca gtacagccac caaggacacc	1440
tacgacgccc ttcacatgca ggccctgccc cctcgctgat aa	1482

<210> 1676  
 <211> 63  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 leader sequence

<400> 1676	
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ccg	63

<210> 1677  
 <211> 744  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> h20A100 scFV

<400> 1677	
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agctgtgccg ccagcggcctt caccttttagc acatacgcca tgagctgggt ccgacaggcc	120

cctggaaaag gccttgaatg ggttgctct atcggcagag ccggcagcac ctactacagc	180
gattctgtga agggcagatt caccatcagc cgggacaacg ccaagaacag cctgtacctg	240
cagatgaact ccctgagagc cgaggacacc gccgtgtact attgtgccag aggacccatc	300
tacaacgact acgacgagtt cgcctattgg ggccagggca cactgggtcac agtcagctct	360
ggcgggtggcg gaagcggagg cgggtggctcc ggtggcggag gcagcgacat cgtgatgaca	420
cagagccctg atagcctggc cgtgtctctg ggagagagag ccaccatcaa ctgcaagagc	480
agccagagcg tgctgtactc cagcaaccag aagaactacc tggcctggta tcagcagaag	540
cccggccagc ctcctaagct gctgatctac tgggccagca ccagagaaaag cggcgtgccc	600
gatagatttt ctggcagcgg ctctggcacc gacttcaccc tgacaattag ctccctgcag	660
gccgaggatg tggccgtgta ctactgtcac cagtacctga gcagcctgac ctttggcggc	720
ggaacaaagg tggaaatcaa gcga	744

<210> 1678  
 <211> 135  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 hinge region

<400> 1678	
acaacaacc ctgccccag acctcctacc ccagccccta caattgccag ccagcctctg	60
agcctgaggc ccgaggcttg tagacctgct gctggcggag ccgtgcacac cagaggactg	120
gatttcgcct gcgac	135

<210> 1679  
 <211> 63  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 leader sequence

<400> 1679	
atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg	60
ccg	63

<210> 1680  
 <211> 744  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> h20A10N scFV

<400> 1680	
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agctgtgccg ccagcggctt cacctttagc acatacgcca tgagctggat cagacaggcc	120
cctggcaaag gcctggaatg ggtggcgtct attggcagag ccggcagcac ctactacagc	180



gactctgtga agggcagatt caccatcagc cgggacaacg ccaagaacag cctgtacctg	240
cagatgaact ccctgagagc cgaggacacc gccgtgtact attgtgccag aggacccatc	300
tacaacgact acgacgagtt cgcctattgg ggccagggca cactgggtcac agtttctagc	360
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cagagccccg ccacactgtc actttctcca ggcgaaagag ccacactgag ctgcaagagc	480
agccagagcg tgctgtactc cagcaaccag aagaactacc tggcctggta tcagcagaag	540
cccggccaag ctctctggct gctgatctat tgggccagca caagagagag cggcatccct	600
gccagatttt ctggcagcgg ctctggcacc gatttcaccc tgaccataag cagcctggaa	660
cctgaggact tcgccgtgta ttactgccac cagtacctga gcagcctgac ctttggcgga	720
ggcaccaagg tggaaatcaa gcgg	744

<210> 1681  
 <211> 135  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 hinge region

<400> 1681	
acaacaaccc ctgccccag acctcctacc ccagccccta caattgccag ccagcctctg	60
agcctgaggc ccgaggcttg tagacctgct gctggcggag ccgtgcacac cagaggactg	120
gatttcgcct gcgac	135

<210> 1682  
 <211> 72  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 transmembrane domain

<400> 1682	
atctacatct gggcgccctt ggccgggact tgtgggggtcc ttctcctgtc actggttatc	60
accctttact gc	72

<210> 1683  
 <211> 126  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> 41BB

<400> 1683	
aaacggggca gaaagaaact cctgtatata ttcaaacaac catttatgag accagtacaa	60
actactcaag aggaagatgg ctgtagctgc cgatttcag aagaagaaga aggaggatgt	120
gaactg	126

<210> 1684  
 <211> 342

<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD3

<400> 1684  
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tataacgagc tcaatctagg acgaagagag gagtacgatg ttttggacaa gagacgtggc 120  
cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgtacaat 180  
gaactgcaga aagataagat ggccggaggcc tacagtgaga ttgggatgaa aggcgagcgc 240  
cggagggggca aggggcacga tggcctttac cagggtctca gtacagccac caaggacacc 300  
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<210> 1685  
<211> 1482  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> pCDH MSCV h20A10-C2 CAR 41BB

<400> 1685  
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ccggaggtgc agctggttga atctggcggc ggacttgtga agcctggcgg atctctgaga 120  
ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg ggtccgacag 180  
gccccctggaa aaggccttga atgggttgcc tctatcggca gagccggcag cacctactac 240  
agcgattctg tgaagggcag attcaccatc agccgggaca acgccaagaa cagcctgtac 300  
ctgcagatga actccctgag agccgaggac accgccgtgt actattgtgc cagaggaccc 360  
atctacaacg actacgacga gttcgcctat tggggccagg gcacactggt cacagtcagc 420  
tctggcgggtg gcggaagcgg aggcgggtggc tccggtggcg gaggcagcga catttgtctg 480  
accagtcctc cagcctcctt ggccgtgtct ccaggacaga gggccaccat cacctgcaag 540  
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag 600  
aaaccaggac aacctcctaa actcctgatt tactgggcca gcaccagaga aagcggggtc 660  
ccagccaggt tcagcggcag tgggtctggg accgatttca ccctcacaat taatcctgtg 720  
gaagctaatt atactgcaaa ttattactgt caccagtacc tgagcagcct gaccttcggc 780  
ggagggacca aggtggagat caaacgaaca acaaccctg ccccagacc tcctacccca 840  
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ggcggagccg tgacaccag aggactggat ttgcctgcg acatctacat ctgggcgccc 960  
ttggccggga cttgtggggt ctttctcctg tctactggtta tcacccttta ctgcaaacgg 1020  
ggcagaaaga aactcctgta tatattcaaa caaccattta tgagaccagt acaaactact 1080  
caagaggaag atggctgtag ctgccgattt ccagaagaag aagaaggagg atgtgaactg 1140  
agagtgaagt tcagcaggag cgcagacgcc cccgcgtaca agcagggcca gaaccagctc 1200

tataacgagc tcaatctagg acgaagagag gagtacgatg ttttggacaa gagacgtggc	1260
cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgtacaat	1320
gaactgcaga aagataagat ggcgagggcc tacagtgaga ttgggatgaa aggcgagcgc	1380
cggaggggca aggggcacga tggcctttac caggtctca gtacagccac caaggacacc	1440
tacgacgccc ttcacatgca ggccctgccc cctcgtgat aa	1482

<210> 1686  
 <211> 63  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 leader sequence

<400> 1686	
atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg	60
ccg	63

<210> 1687  
 <211> 744  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> h20A10C2 scFV

<400> 1687	
gaggtgcagc tggttgaatc tggcggcgga cttgtgaagc ctggcggatc tctgagactg	60
agctgtgccg ccagcggcct cacccttagc acatacgcca tgagctgggt ccgacaggcc	120
cctggaaaag gccttgaatg ggttgccctc atcggcagag ccggcagcac ctactacagc	180
gattctgtga agggcagatt caccatcagc cgggacaacg ccaagaacag cctgtacctg	240
cagatgaact ccctgagagc cgaggacacc gccgtgtact attgtgccag aggacccatc	300
tacaacgact acgacgagtt cgcctattgg ggccagggca cactggtcac agtcagctct	360
ggcgggtggcg gaagcggagg cggctggctcc ggtggcggag gcagcgacat tgtgctgacc	420
cagtctccag cctccttggc cgtgtctcca ggacagaggg ccaccatcac ctgcaagagc	480
agccagagcg tgctgtactc cagcaaccag aagaactacc tggcctggta tcagcagaaa	540
ccaggacaac ctccctaaact cctgatttac tgggccagca ccagagaaaag cggggtccca	600
gccaggttca gcggcagtggt gtctgggacc gatttcaccc tcacaattaa tcctgtggaa	660
gctaatagata ctgcaattaa ttactgtcac cagtacctga gcagcctgac cttcggcgga	720
gggaccaagg tggagatcaa acga	744

<210> 1688  
 <211> 135  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 hinge region

<400> 1688  
acaacaaccc ctgccccag acctcctacc ccagccccta caattgccag ccagcctctg 60  
agcctgaggc ccgaggcttg tagacctgct gctggcggag ccgtgcacac cagaggactg 120  
gatttcgcct gcgac 135

<210> 1689  
<211> 72  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 transmembrane region

<400> 1689  
atctacatct gggcgccctt ggccgggact tgtggggtcc ttctcctgtc actggttata 60  
accctttact gc 72

<210> 1690  
<211> 126  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> 41BB

<400> 1690  
aaacggggca gaaagaaact cctgtatata ttcaaacaac catttatgag accagtacaa 60  
actactcaag aggaagatgg ctgtagctgc cgatttccag aagaagaaga aggaggatgt 120  
gaactg 126

<210> 1691  
<211> 342  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD3

<400> 1691  
agagtgaagt tcagcaggag cgcagacgcc cccgcgtaca agcagggccca gaaccagctc 60  
tataacgagc tcaatctagg acgaagagag gactacgatg ttttggacaa gagacgtggc 120  
cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgtacaat 180  
gaactgcaga aagataagat ggcggaggcc tacagtgaga ttgggatgaa aggcgagcgc 240  
cggagggggca aggggcacga tggcctttac cagggtctca gtacagccac caaggacacc 300  
tacgacgccc ttcacatgca ggccctgccc cctcgctgat aa 342

<210> 1692  
<211> 1479  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> pCDH MSCV h20A10-O CAR CD28 1XX

<400> 1692

atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg	60
ccggagggtgc agctggttga atctggcggc ggacttgtga agcctggcgg atctctgaga	120
ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg ggtccgacag	180
gccccctggaa aaggccttga atgggttgcc tctatcgga gagccggcag cacctactac	240
agcgattctg tgaagggcag attcaccatc agccgggaca acgccaagaa cagcctgtac	300
ctgcagatga actccctgag agccgaggac accgccgtgt actattgtgc cagaggaccc	360
atctacaacg actacgacga gttcgcttat tggggccagg gcacactggt cacagtcagc	420
tctggcgggtg gcggaagcgg aggcgggtggc tccggtggcg gaggcagcga catcgtgatg	480
acacagagcc ctgatagcct ggccgtgtct ctgggagaga gagccaccat caactgcaag	540
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag	600
aagcccggcc agcctcctaa gctgctgata tactgggccg gcaccagaga aagcggcgtg	660
cccgatagat tttctggcag cggtcttggc accgacttca ccctgacaat tagctccctg	720
caggccgagg atgtggccgt gtactactgt caccagtacc tgagcagcct gacctttggc	780
ggcggaaaca aggtggaaat caagcgaaca acaaccctg cccccagacc tcctaccca	840
gccccataca ttgccagcca gcctctgagc ctgaggcccg aggctttag acctgtgct	900
ggcggagccg tgcacaccag aggactggat ttgcctgcg acatctacat ctgggcgccc	960
ttggccggga cttgtggggt ctttctcctg tctactggtta tcacccttta ctgcaggagt	1020
aagaggagca ggctcctgca cagtgactac atgaacatga ctcctagaag acctgggcct	1080
accagaaagc attaccagcc ctatgcccc ccaacgcgact tcgcagccta tcgctccaga	1140
gtgaagttca gcaggagcgc agacgcccc gcgtacaagc agggccagaa ccagctctat	1200
aacgagctca atctaggacg aagagaggag tacgatgttt tggacaagag acgtggccgg	1260
gaccctgaga tggggggaaa gccgagaagg aagaaccctc aggaaggcct gttcaatgaa	1320
ctgcagaaag ataagatggc ggaggccttc agtgagattg ggatgaaagg cgagcgccgg	1380
aggggcaagg ggcacgatgg ctttttcag ggtctcagta cagccaccaa ggacaccttc	1440
gacgcccttc acatgcaggc cctgccccct cgctgataa	1479

<210> 1693  
 <211> 63  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 leader sequence

<400> 1693	
atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg	60
ccg	63

<210> 1694  
 <211> 744  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> h20A100 scFV

<400> 1694  
 gaggtgcagc tggttgaatc tggcggcggg cttgtgaagc ctggcggatc tctgagactg 60  
 agctgtgccg ccagcggcctt cacccttagc acatacgcca tgagctgggt ccgacaggcc 120  
 cctggaaaag gccttgaatg ggttgccctc atcggcagag ccggcagcac ctactacagc 180  
 gattctgtga agggcagatt caccatcagc cgggacaacg ccaagaacag cctgtacctg 240  
 cagatgaact ccctgagagc cgaggacacc gccgtgtact attgtgccag aggacccatc 300  
 tacaacgact acgacgagtt cgcctattgg ggccagggca cactgggtcac agtcagctct 360  
 ggccggtggcg gaagcggagg cgggtggctcc ggtggcggag gcagcgacat cgtgatgaca 420  
 cagagccctg atagcctggc cgtgtctctg ggagagagag ccaccatcaa ctgcaagagc 480  
 agccagagcg tegtgtactc cagcaaccag aagaactacc tggcctggta tcagcagaag 540  
 cccggccagc ctcctaagct gctgatctac tgggccagca ccagagaaaag cggcgtgccc 600  
 gatagatttt ctggcagcgg ctctggcacc gacttcaccc tgacaattag ctccctgcag 660  
 gccgaggatg tggccgtgta ctactgtcac cagtacctga gcagcctgac ctttggcggc 720  
 ggaacaaagg tggaaatcaa gcga 744

<210> 1695  
 <211> 135  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 hinge region

<400> 1695  
 acaacaaccc ctgccccag acctcctacc ccagccccta caattgccag ccagcctctg 60  
 agcctgaggc ccgaggcttg tagacctgct gctggcggag ccgtgcacac cagaggactg 120  
 gatttcgcct gcgac 135

<210> 1696  
 <211> 72  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 transmembrane region

<400> 1696  
 atctacatct gggcgccctt ggccgggact tgtgggggcc ttctcctgtc actggttata 60  
 accctttact gc 72

<210> 1697  
 <211> 123  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD28

<400> 1697  
aggagtaaga ggagcaggct cctgcacagt gactacatga acatgactcc tagaagacct 60  
gggcctacca gaaagcatta ccagccctat gccccaccac gcgacttcgc agcctatcgc 120  
tcc 123

<210> 1698  
<211> 342  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD3 1XX

<400> 1698  
agagtgaagt tcagcaggag cgcagacgcc cccgcgtaca agcagggcca gaaccagctc 60  
tataacgagc tcaatctagg acgaagagag gagtacgatg ttttggacaa gagacgtggc 120  
cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgttcaat 180  
gaactgcaga aagataagat ggccggaggcc ttcagtgaga ttgggatgaa aggcgagcgc 240  
cggaggggca aggggcacga tggccttttc cagggtctca gtacagccac caaggacacc 300  
ttcgacgccc ttcacatgca ggccctgccc cctcgctgat aa 342

<210> 1699  
<211> 1479  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> pCDH MSCV h20A10-N CAR CD28 1XX

<400> 1699  
atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg 60  
ccgcagggtgc agctggttga atctggcggc ggacttgtga agcctggcgg atctctgaga 120  
ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg gatcagacag 180  
gcccctggca aaggcctgga atgggtggcg tctattggca gagccggcag cacctactac 240  
agcgactctg tgaagggcag attcaccatc agccgggaca acgccaagaa cagcctgtac 300  
ctgcagatga actccctgag agccgaggac accgccgtgt actattgtgc cagaggaccc 360  
atctacaacg actacgacga gttgcctat tggggccagg gcacactggt cacagtttct 420  
agcggcgggtg gcggaagcgg aggcgggtggc tccggtggcg gaggcagcga aattgtgctg 480  
acacagagcc ccgccacact gtcactttct ccaggcgaaa gagccacact gagctgcaag 540  
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag 600  
aagcccggcc aagctcctcg gctgctgatc tattgggcca gcacaagaga gagcggcatc 660  
cctgccagat tttctggcag cggctctggc accgatttca ccctgaccat aagcagcctg 720  
gaacctgagg acttcgccgt gtattactgc caccagtacc tgagcagcct gacctttggc 780  
ggaggcacca aggtggaaat caagcggaca acaaccctg ccccagacc tcctacccca 840  
gcccctacaa ttgccagcca gcctctgagc ctgaggcccc aggctttag acctgctgct 900

ggcggagccg tgcacaccag aggactggat ttcgcctgcg acatctacat ctgggcgccc	960
ttggccggga cttgtggggt ctttctcctg tctactggtta tcacccttta ctgcaggagt	1020
aagaggagca ggctcctgca cagtgactac atgaacatga ctcctagaag acctgggcct	1080
accagaaagc attaccagcc ctatgcccc ccacgcgact tcgcagccta tcgctccaga	1140
gtgaagttca gcaggagcgc agacgcccc gcgtacaagc agggccagaa ccagctctat	1200
aacgagctca atctaggacg aagagaggag tacgatgttt tggacaagag acgtggccgg	1260
gaccctgaga tggggggaaa gccgagaagg aagaaccctc aggaaggcct gttcaatgaa	1320
ctgcagaaag ataagatggc ggaggccttc agtgagattg ggatgaaagg cgagcgccgg	1380
aggggcaagg ggacacgatgg ctttttccag ggtctcagta cagccaccaa ggacaccttc	1440
gacgcccttc acatgcaggc cctgccccct cgctgataa	1479

<210> 1700  
 <211> 63  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 leader sequence

<400> 1700	
atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg	60
ccg	63

<210> 1701  
 <211> 744  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> h20A10N scFV

<400> 1701	
caggtgcagc tggttgaatc tggcggcgga cttgtgaagc ctggcggatc tctgagactg	60
agctgtgccg ccagcggcctt cacccttagc acatacgcca tgagctggat cagacaggcc	120
cctggcaaag gcctggaatg ggtggcgtct attggcagag ccggcagcac ctactacagc	180
gactctgtga agggcagatt caccatcagc cgggacaacg ccaagaacag cctgtacctg	240
cagatgaact ccctgagagc cgaggacacc gccgtgtact attgtgccag aggacccatc	300
tacaacgact acgacgagtt cgcctattgg ggccagggca cactgggtcac agtttctagc	360
ggcgggtggcg gaagcggagg cgggtggctcc ggtggcggag gcagcgaaat tgtgctgaca	420
cagagccccg ccacactgtc actttctcca ggcgaaagag ccacactgag ctgcaagagc	480
agccagagcg tgctgtactc cagcaaccag aagaactacc tggcctggta tcagcagaag	540
cccggccaag ctctctggct gctgatctat tgggccagca caagagagag cggcatccct	600
gccagatttt ctggcagcgg ctctggcacc gatttcaccc tgaccataag cagcctggaa	660
cctgaggact tcgccgtgta ttactgccac cagtacctga gcagcctgac ctttggcgga	720



<210> 1702  
<211> 135  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 hinge region

<400> 1702  
acaacaaccc ctgccccag acctcctacc ccagccccta caattgccag ccagcctctg 60  
agcctgaggc ccgaggcttg tagacctgct gctggcggag ccgtgcacac cagaggactg 120  
gatttcgcct gcgac 135

<210> 1703  
<211> 72  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 transmembrane domain

<400> 1703  
atctacatct gggcgccctt ggccgggact tgtggggtcc ttctcctgtc actggttatc 60  
accctttact gc 72

<210> 1704  
<211> 123  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD28

<400> 1704  
aggagtaaga ggagcaggct cctgcacagt gactacatga acatgactcc tagaagacct 60  
gggcctacca gaaagcatta ccagccctat gccccaccac gcgacttcgc agcctatcgc 120  
tcc 123

<210> 1705  
<211> 342  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD3 1XX

<400> 1705  
agagtgaagt tcagcaggag cgcagacgcc cccgcgtaca agcagggcca gaaccagctc 60  
tataacgagc tcaatctagg acgaagagag gagtacgatg ttttgacaa gagacgtggc 120  
cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgttcaat 180  
gaactgcaga aagataagat ggcggaggcc ttcagtgaga ttgggatgaa aggcgagcgc 240  
cggaggggca aggggcacga tggccttttc cagggtctca gtacagccac caaggacacc 300  
ttcgagccc ttcacatgca ggccctgccc cctcgctgat aa 342

<210> 1706  
<211> 1479  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> pCDH MSCV h20A10-C2 CAR CD28 1XX

<400> 1706  
atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg 60  
ccggagggtgc agctgggtga atctggcggc ggacttgtag agcctggcgg atctctgaga 120  
ctgagctgtg ccgccagcgg cttcaccttt agcacatacg ccatgagctg ggtccgacag 180  
gcccctggaa aaggccttga atgggttgcc tctatcggca gagccggcag cacctactac 240  
agcgattctg tgaagggcag attcaccatc agccgggaca acgccaagaa cagcctgtac 300  
ctgcagatga actccctgag agccgaggac accgccgtgt actattgtgc cagaggaccc 360  
atctacaacg actacgacga gttcgcctat tggggccagg gcacactggt cacagtcagc 420  
tctggcgggtg gcggaagcgg aggcgggtggc tccggtggcg gaggcagcga cattgtgctg 480  
accagttctc cagcctcctt ggccgtgtct ccaggacaga gggccacat cacctgcaag 540  
agcagccaga gcgtgctgta ctccagcaac cagaagaact acctggcctg gtatcagcag 600  
aaaccaggac aacctcctaa actcctgatt tactgggcca gcaccagaga aagcggggtc 660  
ccagccaggt tcagcggcag tgggtctggg accgatttca ccctcacaat taatcctgtg 720  
gaagctaatt atactgcaaa ttattactgt caccagtacc tgagcagcct gaccttcggc 780  
ggagggacca agtgaggat caaacgaaca acaaccctg ccccgagacc tcctacccca 840  
gcccctacaa ttgccagcca gcctctgagc ctgaggcccg aggcctttag acctgctgct 900  
ggcggagccg tgcacaccag aggactggat ttgcctgctg acatctacat ctgggcgccc 960  
ttggccggga cttgtgggt cttctcctg tcaactgtta tcacccttta ctgcaggagt 1020  
aagaggagca ggctcctgca cagtgactac atgaacatga ctctagaag acctgggcct 1080  
accagaaagc attaccagcc ctatgcccc cccgcgact tcgcagccta tcgctccaga 1140  
gtgaagtcca gcaggagcgc agacgcccc gcgtacaagc agggccagaa ccagctctat 1200  
aacgagctca atctaggacg aagagaggag tacgatgttt tggacaagag acgtggccgg 1260  
gacctgaga tggggggaaa gccgagaagg aagaaccctc aggaaggcct gttcaatgaa 1320  
ctgcagaaag ataagatggc ggaggccttc agtgagattg ggatgaaagg cgagcgccgg 1380  
aggggcaagg ggcacgatgg cttttccag ggtctcagta cagccaccaa ggacaccttc 1440  
gacgcccttc acatgcaggc cctgccccct cgctgataa 1479

<210> 1707  
<211> 63  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 leader sequence

<400> 1707  
atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg 60  
ccg 63

<210> 1708  
<211> 744  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> h20A10C2 scFV

<400> 1708  
gaggtgcagc tggttgaatc tggcggcgga cttgtgaagc ctggcggatc tctgagactg 60  
agctgtgccg ccagcggcctt caccttttagc acatacgcca tgagctgggt ccgacaggcc 120  
cctggaaaag gccttgaatg ggttgccctt atcggcagag ccggcagcac ctactacagc 180  
gattctgtga agggcagatt caccatcagc cgggacaacg ccaagaacag cctgtacctg 240  
cagatgaact ccctgagagc cgaggacacc gccgtgtact attgtgccag aggacccatc 300  
tacaacgact acgacgagtt cgcctattgg ggccagggca cactgggtcac agtcagctct 360  
ggcgggtggcg gaagcggagg cgggtggctcc ggtggcggag gcagcgacat tgtgctgacc 420  
cagtctccag cctccttggc cgtgtctcca ggacagaggg ccaccatcac ctgcaagagc 480  
agccagagcg tgctgtactc cagcaaccag aagaactacc tggcctggta tcagcagaaa 540  
ccaggacaac ctctaaact cctgatttac tgggccagca ccagagaaaag cgggggtcca 600  
gccaggttca gcggcagtg gttctgggacc gatttcaccc tcacaattaa tcctgtggaa 660  
gctaatagata ctgcaaatta ttactgtcac cagtacctga gcagcctgac cttcggcgga 720  
gggaccaagg tggagatcaa acga 744

<210> 1709  
<211> 135  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 hinge region

<400> 1709  
acaacaaccc ctgccccag acctcctacc ccagccccta caattgccag ccagcctctg 60  
agcctgaggc ccgaggcttg tagacctgct gctggcggag ccgtgcacac cagaggactg 120  
gatttcgcct gcgac 135

<210> 1710  
<211> 72  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 transmembrane region

<400> 1710  
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<210> 1711  
 <211> 123  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD28

<400> 1711  
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 gggcctacca gaaagcatta ccagccctat gccccaccac gcgacttcgc agcctatcgc 120  
 tcc 123

<210> 1712  
 <211> 342  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD3 1XX

<400> 1712  
 agagtgaagt tcagcaggag cgcagacgcc cccgcgtaca agcagggcca gaaccagctc 60  
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 cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgttcaat 180  
 gaactgcaga aagataagat ggccggaggcc ttcagtgaga ttgggatgaa aggcgagcgc 240  
 cggaggggca aggggcacga tggccttttc cagggtctca gtacagccac caaggacacc 300  
 ttcgacgccc ttcacatgca ggccctgccc cctcgctgat aa 342

<210> 1713  
 <211> 2380  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> pCDH MSCV hC2 CAR 41BB 6xNFATFoxP3 IL18

<400> 1713  
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 ctctcctgtg cagcctctgg attcaccttc agtggctatg ccatgagctg ggtccgccag 180  
 gctccaggga aggggctgga gtgggtctca accattagta gtggcggaac ctacatatac 240  
 taccgact cagtgaaggg ccgattcacc atctccagag acaacgcaa gaactcactg 300  
 tatctgcaaa tgaacagcct gagagccgag gacacggccg tgtattactg tgcgagactt 360  
 ggggggggata attactacga atacttcgat gtctggggca aagggaccac ggtcaccgtc 420  
 tcctccggcg gtggcggtgc cggcggtggc ggatccggcg gtggcggtgc cgacattgtg 480  
 ctgaccagct ctccagcctc cttggccgtg tctccaggac agagggccac catcacctgc 540

agagccagta agagtgtcag taccagcgga tactcctaca tgcactggta tcagcagaaa	600
ccaggacaac ctccctaaact cctgattttac ctggcatcca atctggagag cgggggtcca	660
gccaggttca gcggcagtgg gtctgggacc gatttcaccc tcacaattaa tcctgtggaa	720
gctaatagata ctgcaaatta ttactgtcag cacagtaggg agctgccttt cacattcggc	780
ggaggggacca aggtggagat caaacgaact acaacaaccc ctgccccag acctcctacc	840
ccagccccta caattgccag ccagcctctg agcctgaggc ccgaggcttg tagacctgct	900
gctggcggag ccgtgcacac cagaggactg gatttcgcct gcgacatcta catctgggcg	960
cccttggccg ggacttgtgg ggtccttctc ctgtcactgg ttatcacctt ttactgcaaa	1020
cggggcagaa agaaactcct gtatatattc aaacaacat ttatgagacc agtacaaact	1080
actcaagagg aagatggctg tagctgccga tttccagaag aagaagaagg aggatgtgaa	1140
ctgagagtga agttcagcag gagcgagac gccccgcgt acaagcaggg ccagaaccag	1200
ctctataacg agctcaatct aggacgaaga gaggagtacg atgttttgga caagagacgt	1260
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aatgaactgc agaaagataa gatggcggag gcctacagt agattgggat gaaaggcgag	1380
cgccggaggg gcaaggggca cgatggcctt taccagggtc tcagtacagc caccaaggac	1440
acctacgacg cccttcacat gcaggccctg cccctcgt gataagtta aactgccaga	1500
acattttctt ggccctaactg gccggtaccg gcttcatttt ttccatttac tgcagaggct	1560
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gtgaaccgtc agatcgctg gagacgcat ccacgctgtt ttgacctca tagaagacac	1800
cgggaccgat ccagcctcga gagacccaat gctagccacc atggccttac cagtgaccgc	1860
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cagacccctg ttcgaggaca tgaccgacag cgactgcaga gacaacgcc ctcggaccat	2040
cttcatcatc agcatgtaca aggacagcca gcctagaggc atggccgtga ccatctctgt	2100
gaagtgcgag aagatcagca ccctgagctg cgagaacaag atcatcagct tcaaagagat	2160
gaaccgcgcg gacaacatca aggacaccaa gagcgacatc atattcttcc agcggagcgt	2220
gcccggccac gacaacaaga tgcagtttga gagcagcagc tacgagggt acttcctggc	2280
ctgcgagaaa gagcgggacc tgttcaagct gatcctgaag aaagaggacg aactgggcga	2340
ccgcagcatc atgttcaccg tgcagaacga ggactgataa	2380

<210> 1714

<211> 63

<212> DNA

<213> Artificial Sequence

<220>

<223> CD8 leader sequence

<400> 1714

atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg 60  
ccg 63

<210> 1715

<211> 747

<212> DNA

<213> Artificial Sequence

<220>

<223> C2 scFV

<400> 1715

gagggtgcagc tgggtggagtc tgggggaggc ctggtcaagc ctgggggggtc cctgagactc 60  
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ccagggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac 180  
cccgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240  
ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagacttggg 300  
ggggataatt actacgaata cttcgatgtc tggggcaaag ggaccacggt caccgtctcc 360  
tccggcgggtg gcggatccgg cgggtggcga tccggcgggt gcggatccga cattgtgctg 420  
accagtctc cagcctcctt ggccgtgtct ccaggacaga gggccaccat cacctgcaga 480  
gccagtaaga gtgtcagtac cagcggatac tcctacatgc actggtatca gcagaaacca 540  
ggacaacctc ctaaactcct gatttacctg gcatccaatc tggagagcgg ggtcccagcc 600  
aggttcagcg gcagtgggtc tgggaccgat ttcaccctca caattaatcc tgtggaagct 660  
aatgatactg caaattatta ctgtcagcac agtagggagc tgcctttcac attcggcgga 720  
gggaccaagg tggagatcaa acgaact 747

<210> 1716

<211> 135

<212> DNA

<213> Artificial Sequence

<220>

<223> CD8 hinge region

<400> 1716

acaacaaccc ctgccccag acctcctacc ccagccccta caattgccag ccagcctctg 60  
agcctgaggc ccgaggcttg tagacctgct gctggcggag ccgtgcacac cagaggactg 120  
gatttcgcct gcgac 135

<210> 1717

<211> 72

<212> DNA

<213> Artificial Sequence

<220>

<223> CD8 transmembrane region

<400> 1717

atctacatct gggcgccctt ggccgggact tgtgggggcc ttctcctgtc actggttatc 60

accctttact gc 72

<210> 1718

<211> 126

<212> DNA

<213> Artificial Sequence

<220>

<223> 41BB

<400> 1718

aaacggggca gaaagaaact cctgtatata ttcaaacaac catttatgag accagtacaa 60

actactcaag aggaagatgg ctgtagctgc cgatttccag aagaagaaga aggaggatgt 120

gaactg 126

<210> 1719

<211> 336

<212> DNA

<213> Artificial Sequence

<220>

<223> CD3

<400> 1719

agagtgaagt tcagcaggag cgcagacgcc cccgcgtaca agcagggccca gaaccagctc 60

tataacgagc tcaatctagg acgaagagag gagtacgatg ttttggacaa gagacgtggc 120

cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgtacaat 180

gaactgcaga aagataagat ggccggaggcc tacagtgaga ttgggatgaa aggcgagcgc 240

cggaggggca aggggcacga tggcctttac cagggtctca gtacagccac caaggacacc 300

tacgacgccc ttcacatgca ggccctgccc cctcgc 336

<210> 1720

<211> 50

<212> DNA

<213> Artificial Sequence

<220>

<223> plasmid

<400> 1720

tgataagttt aaactgccag aacatttctc tggcctaact ggccggtacc 50

<210> 1721

<211> 168

<212> DNA

<213> Artificial Sequence

<220>

<223> 6xNFAT FoxP3

<400> 1721

ggcttcattt tttccattta ctgcagaggc ttcatTTTTT ccattttactg cagaggcttc 60

attttttcca tttactgcag aggccttcatt ttttccattt actgcagagg cttcatTTTT 120

tccattttact gcagaggctt cattttttcc attttactgca gaactagt 168

<210> 1722  
 <211> 118  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> mCMV  
  
 <400> 1722  
 taggcgtgta cggtagggagg cctatataag cagagctcgt ttagtgaacc gtcagatcgc 60  
 ctggagacgc catccacgct gttttgacct ccatagaaga caccgggacc gatccagc 118  
  
 <210> 1723  
 <211> 25  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> plasmid  
  
 <400> 1723  
 ctcgagagac ccaatgctag ccacc 25  
  
 <210> 1724  
 <211> 63  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> CD8 leader sequence  
  
 <400> 1724  
 atggccttac cagtgaccgc cttgctcctg ccgctggcct tgctgctcca cgccgccagg 60  
 ccg 63  
  
 <210> 1725  
 <211> 477  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> IL18  
  
 <400> 1725  
 tacttcggca agctggaaag caagctgagc gtgatccgga acctgaacga ccaggtgctg 60  
 ttcacatgatc agggcaacag acccctgttc gaggacatga ccgacagcga ctgcagagac 120  
 aacgcccctc ggaccatctt catcatcagc atgtacaagg acagccagcc tagaggcatg 180  
 gccgtgacca tctctgtgaa gtgcgagaag atcagcaccc tgagctgcga gaacaagatc 240  
 atcagcttca aagagatgaa cccgccggac aacatcaagg acaccaagag cgacatcata 300  
 ttcttccagc ggagcgtgcc cggccacgac aacaagatgc agtttgagag cagcagctac 360  
 gagggctact tcctggcctg cgagaaagag cgggacctgt tcaagctgat cctgaagaaa 420  
 gaggacgaac tgggcgaccg cagcatcatg ttcaccgtgc agaacgagga ctgataa 477  
  
 <210> 1726



<211> 1482  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> pCDH MSCV hC2 CAR CD28 1XX

<400> 1726  
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ctctcctgtg cagcctctgg attcaccttc agtggctatg ccatgagctg ggtccgccag 180  
gctccaggga aggggctgga gtgggtctca accattagta gtggcggaac ctacatatac 240  
taccgccact cagtgaaggg ccgattcacc atctccagag acaacgcaa gaactcactg 300  
tatctgcaaa tgaacagcct gagagccgag gacacggcgg tgtattactg tgcgagactt 360  
gggggggata attactacga atacttcgat gtctggggca aagggaccac ggtcaccgtc 420  
tcctccggcg gtggcggatc cggcgggtggc ggatccggcg gtggcggatc cgacattgtg 480  
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agagccagta agagtgtcag taccagcgga tactcctaca tgcactggta tcagcagaaa 600  
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gccaggttca gcggcagtggt gtctgggacc gatttcaccc tcacaattaa tcctgtggaa 720  
gctaatagata ctgcaaatta ttactgtcag cacagtaggg agctgccttt cacattcggc 780  
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gaactgcaga aagataagat ggccggaggcc ttcagtgaga ttgggatgaa aggcgagcgc 1380  
cggaggggca aggggcacga tggccttttc cagggtctca gtacagccac caaggacacc 1440  
ttcgacgccc ttcacatgca ggccctgccc cctcgctgat aa 1482

<210> 1727  
<211> 63  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 leader sequence

<400> 1727  
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<210> 1728  
 <211> 747  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> C2 scFV

<400> 1728  
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 tcctgtgcag cctctggatt caccttcagt ggctatgcc a tgagctgggt ccgccaggct 120  
 ccagggaagg ggctggagtg ggtctcaacc attagtagtg gcggaaccta catatactac 180  
 cccgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240  
 ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagacttggg 300  
 ggggataatt actacgaata cttcgatgtc tggggcaaag ggaccacggt caccgtctcc 360  
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 ggacaacctc ctaaactcct gatttacctg gcattccaatc tggagagcgg ggtcccagcc 600  
 aggttcagcg gcagtgggtc tgggaccgat ttcaccctca caattaatcc tgtggaagct 660  
 aatgatactg caaattatta ctgtcagcac agtagggagc tgcctttcac attcggcgga 720  
 gggaccaagg tggagatcaa acgaact 747

<210> 1729  
 <211> 135  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 hinge region

<400> 1729  
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 gatttcgcct gcgac 135

<210> 1730  
 <211> 72  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> CD8 transmembrane region

<400> 1730  
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 accctttact gc 72

<210> 1731  
 <211> 123  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> CD28  
  
 <400> 1731  
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 gggcctacca gaaagcatta ccagccctat gccccaccac gcgacttcgc agcctatcgc 120  
 tcc 123

<210> 1732  
 <211> 342  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> CD3 1XX  
  
 <400> 1732  
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 cgggaccctg agatgggggg aaagccgaga aggaagaacc ctcaggaagg cctgttcaat 180  
 gaactgcaga aagataagat ggcggaggcc ttcagtgaga ttgggatgaa aggcgagcgc 240  
 cggaggggca aggggcacga tggccttttc cagggtctca gtacagccac caaggacacc 300  
 ttcgagccc ttcacatgca ggccctgccc cctcgctgat aa 342

<210> 1733  
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 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B12 light chain variable framework 2 (FR2) sequence  
  
 <400> 1733  
 tggatcagc agaaaccagg acagccaccc aaactcctca tctat 45

<210> 1734  
 <211> 15  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B12 light chain variable framework 2 (FR2) sequence  
  
 <400> 1734

Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Ile Tyr  
 1 5 10 15

<210> 1735  
 <211> 21  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse B12 light chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 1735  
 cttgcatcca ccctagattc t 21

<210> 1736  
 <211> 7  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse B12 light chain variable complementarity determining regions 2 (CDR2) sequence  
  
 <400> 1736  
  
 Leu Ala Ser Thr Leu Asp Ser  
 1 5

<210> 1737  
 <211> 96  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse B12 light chain variable framework 3 (FR3) sequence  
  
 <400> 1737  
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 cctgtggagg ctgatgatgc tgcaacctat tactgt 96

<210> 1738  
 <211> 32  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> Mouse B12 light chain variable framework 3 (FR3) sequence  
  
 <400> 1738  
  
 Gly Val Pro Ala Arg Phe Ser Gly Ser Gly Ser Arg Thr Asp Phe Thr  
 1 5 10 15

Leu Thr Ile Asp Pro Val Glu Ala Asp Asp Ala Ala Thr Tyr Tyr Cys  
 20 25 30

<210> 1739  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Mouse B12 light chain variable complementarity determining regions 3 (CDR3) sequence  
  
 <400> 1739  
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<210> 1740  
 <211> 9  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B12 light chain variable complementarity determining regions 3 (CDR3) sequence  
  
 <400> 1740

Gln Gln Asn Asn Glu Asp Pro Pro Thr  
 1 5

<210> 1741  
 <211> 31  
 <212> DNA  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B12 light chain variable framework 4 (FR4) sequence  
  
 <400> 1741  
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31

<210> 1742  
 <211> 10  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> Mouse B12 light chain variable framework 4 (FR4) sequence  
  
 <400> 1742

Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys  
 1 5 10

<210> 1743  
 <211> 10  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> epitope to which NME1 and NME7AB bind part of the 10 membrane proximal PRT  
  
 <400> 1743

Pro Phe Pro Phe Ser Ala Gln Ser Gly Ala  
 1 5 10

<210> 1744  
 <211> 11  
 <212> PRT  
 <213> Artificial Sequence  
  
 <220>  
 <223> epitope to which NME1 and NME7AB bind part of the 10 membrane proximal PRT  
  
 <400> 1744

Ser Asn Ile Lys Phe Arg Pro Gly Ser Val Val  
 1 5 10

<210> 1745  
<211> 7  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> epitope to which NME1 and NME7AB bind part of the 10 membrane proximal PRT

<400> 1745

Ala Ser Arg Tyr Asn Leu Thr  
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<210> 1746  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Fragment of PSMGFR

<400> 1746

Gly Thr Ile Asn Val His Asp Val Glu Thr  
1 5 10

<210> 1747  
<211> 4  
<212> PRT  
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<220>  
<223> Fragment of PSMGFR

<400> 1747

Phe Pro Phe Ser  
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<210> 1748  
<211> 20  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Fragment of PSMGFR

<400> 1748

Ser Asn Ile Lys Phe Arg Pro Gly Ser Val Val Val Gln Leu Thr Leu  
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Ala Phe Arg Glu  
20

<210> 1749  
<211> 9  
<212> PRT  
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<220>

<223> Fragment of PSMGFR

<400> 1749

Gln Phe Asn Gln Tyr Lys Thr Glu Ala  
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<210> 1750

<211> 9

<212> PRT

<213> Artificial Sequence

<220>

<223> Fragment of PSMGFR

<400> 1750

Val Gln Leu Thr Leu Ala Phe Arg Glu  
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<210> 1751

<211> 5

<212> PRT

<213> Artificial Sequence

<220>

<223> Fragment of PSMGFR

<400> 1751

Ser Val Ser Asp Val  
1 5

<210> 1752

<211> 627

<212> DNA

<213> Artificial Sequence

<220>

<223> IL18 sequence

<400> 1752

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ggcaagctgg aaagcaagct gagcgtgatc cggaacctga acgaccaggt gctgttcac	180
gatcaggga acagaccctt gttcgaggac atgaccgaca gcgactgcag agacaacgcc	240
cctcggacca tcttcacat cagcatgtac aaggacagcc agcctagagg catggccgtg	300
accatctctg tgaagtgcga gaagatcagc accctgagct gcgagaacaa gatcatcagc	360
ttcaaagaga tgaacccgcc ggacaacatc aaggacacca agagcgacat catattcttc	420
cagcggagcg tgcccgccca cgacaacaag atgcagtttg agagcagcag ctacgagggc	480
tacttcctgg cctgcgagaa agagcgggac ctgttcaagc tgatcctgaa gaaagaggac	540
gaactgggag accgcagcat catgttcacc gtgcagaacg aggactgata aaagcttggc	600
aatccggtac tgttggtaaa gccacca	627

<210> 1753

<211> 207  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> IL18 sequence

<400> 1753

Ile Glu Asp Thr Gly Thr Asp Pro Ala Ser Arg Asp Pro Met Leu Ala  
1 5 10 15

Thr Met Ala Leu Pro Val Thr Ala Leu Leu Pro Leu Ala Leu Leu  
20 25 30

Leu His Ala Ala Arg Pro Tyr Phe Gly Lys Leu Glu Ser Lys Leu Ser  
35 40 45

Val Ile Arg Asn Leu Asn Asp Gln Val Leu Phe Ile Asp Gln Gly Asn  
50 55 60

Arg Pro Leu Phe Glu Asp Met Thr Asp Ser Asp Cys Arg Asp Asn Ala  
65 70 75 80

Pro Arg Thr Ile Phe Ile Ile Ser Met Tyr Lys Asp Ser Gln Pro Arg  
85 90 95

Gly Met Ala Val Thr Ile Ser Val Lys Cys Glu Lys Ile Ser Thr Leu  
100 105 110

Ser Cys Glu Asn Lys Ile Ile Ser Phe Lys Glu Met Asn Pro Pro Asp  
115 120 125

Asn Ile Lys Asp Thr Lys Ser Asp Ile Ile Phe Phe Gln Arg Ser Val  
130 135 140

Pro Gly His Asp Asn Lys Met Gln Phe Glu Ser Ser Ser Tyr Glu Gly  
145 150 155 160

Tyr Phe Leu Ala Cys Glu Lys Glu Arg Asp Leu Phe Lys Leu Ile Leu  
165 170 175

Lys Lys Glu Asp Glu Leu Gly Asp Arg Ser Ile Met Phe Thr Val Gln  
180 185 190

Asn Glu Asp Lys Leu Gly Asn Pro Val Leu Leu Val Lys Pro Pro  
195 200 205

<210> 1754  
<211> 56  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Primer



<400> 1754  
agggagaccc aagctggcta gttaagcttg gatggcctta ccagtgaccg ccttgc 56

<210> 1755  
<211> 48  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Primer

<400> 1755  
taggccagag aaatgttctg gcattatcag cgagggggca gggcctgc 48

<210> 1756  
<211> 19  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Primer

<400> 1756  
tgccagaaca tttctctgg 19

<210> 1757  
<211> 52  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Primer

<400> 1757  
acagtcgagg ctgatcagcg ggtttaaact tatcagtcct cgttctgcac gg 52

<210> 1758  
<211> 60  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Primer

<400> 1758  
atgcaggccc tgccccctcg ctgataagtt taaactgcc aacatttct ctggcctaac 60

<210> 1759  
<211> 59  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Primer

<400> 1759  
accggagcga tcgcagatcc ttcgcggccg cttatcagtc ctcgttctgc acggtgaac 59

<210> 1760  
<211> 31  
<212> DNA  
<213> Artificial Sequence

<220>		
<223>	Primer	
<400>	1760	
attgcactag ttgaaagacc ccacctgtag g		31
<210>	1761	
<211>	26	
<212>	DNA	
<213>	Artificial Sequence	
<220>		
<223>	Primer	
<400>	1761	
aatgctctag aatacgggta tccagg		26
<210>	1762	
<211>	30	
<212>	DNA	
<213>	Artificial Sequence	
<220>		
<223>	Primer	
<400>	1762	
atagcgaatt cgtaccgagg gccaccatgg		30
<210>	1763	
<211>	44	
<212>	DNA	
<213>	Artificial Sequence	
<220>		
<223>	Primer	
<400>	1763	
taggcctccc accgtacacg cctaggtacc acgccttctg tatg		44
<210>	1764	
<211>	44	
<212>	DNA	
<213>	Artificial Sequence	
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<223>	Primer	
<400>	1764	
taggcctccc accgtacacg cctaggtacc tctgcagtaa atgg		44
<210>	1765	
<211>	19	
<212>	DNA	
<213>	Artificial Sequence	
<220>		
<223>	Primer	
<400>	1765	
taaggccatg gtggctagc		19
<210>	1766	
<211>	32	

<212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Primer

<400> 1766  
 aataagttta aactgccaga acatttctct gg 32

<210> 1767  
 <211> 36  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Primer

<400> 1767  
 atatagcggc cgcttatcag tcctcgttct gcacgg 36

<210> 1768  
 <211> 347  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> 6x FoxP3NFAT mCMV

<400> 1768  
 agaacatttc tctggcctaa ctggccggta ccggcttcat tttttccatt tactgcagag 60  
 gcttcatttt ttccatttac tgcagaggct tcattttttc catttactgc agaggcttca 120  
 ttttttccat ttactgcaga ggcttcattt ttccattta ctgcagaggc ttcatttttt 180  
 ccatttactg cagaactagt taggcgtgta cgggtgggagg cctatataag cagagctcgt 240  
 ttagtgaacc gtcagatcgc ctggagacgc catccacgct gttttgacct ccatagaaga 300  
 caccgggacc gatccagcct cgagagaccc aatgctagcc accatgg 347

<210> 1769  
 <211> 116  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> 6x FoxP3NFAT mCMV

<400> 1769

Arg Thr Phe Leu Trp Pro Asn Trp Pro Val Pro Ala Ser Phe Phe Pro  
 1 5 10 15

Phe Thr Ala Glu Ala Ser Phe Phe Pro Phe Thr Ala Glu Ala Ser Phe  
 20 25 30

Phe Pro Phe Thr Ala Glu Ala Ser Phe Phe Pro Phe Thr Ala Glu Ala  
 35 40 45

Ser Phe Phe Pro Phe Thr Ala Glu Ala Ser Phe Phe Pro Phe Thr Ala  
 50 55 60

Glu Leu Val Arg Arg Val Arg Trp Glu Ala Tyr Ile Ser Arg Ala Arg  
65 70 75 80

Leu Val Asn Arg Gln Ile Ala Trp Arg Arg His Pro Arg Cys Phe Asp  
85 90 95

Leu His Arg Arg His Arg Asp Arg Ser Ser Leu Glu Arg Pro Asn Ala  
100 105 110

Ser His His Gly  
115

<210> 1770  
<211> 343  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> 6x FoxP3NFAT mIL2P

<400> 1770  
agaacatttc tctggcctaa ctggccggta ccggcttcat tttttccatt tactgcagag 60  
gcttcatttt ttccatttac tgcagaggct tcattttttc catttactgc agaggcttca 120  
ttttttccat ttactgcaga ggcttcattt tttccattta ctgcagaggc ttcatttttt 180  
ccatttactg cagaactagt cattttgaca cccccataat atttttccag aattaacagt 240  
ataaaattgca tctcttgttc aagagttccc tatcactctc tttaatcact actcacagta 300  
acctcaactc ctgcctcgag agaccaatg ctagccacca tgg 343

<210> 1771  
<211> 113  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> 6x FoxP3NFAT mIL2P

<220>  
<221> misc\_feature  
<222> (113)..(113)  
<223> Xaa can be any naturally occurring amino acid

<400> 1771

Arg Thr Phe Leu Trp Pro Asn Trp Pro Val Pro Ala Ser Phe Phe Pro  
1 5 10 15

Phe Thr Ala Glu Ala Ser Phe Phe Pro Phe Thr Ala Glu Ala Ser Phe  
20 25 30

Phe Pro Phe Thr Ala Glu Ala Ser Phe Phe Pro Phe Thr Ala Glu Ala  
35 40 45

Ser Phe Phe Pro Phe Thr Ala Glu Ala Ser Phe Phe Pro Phe Thr Ala  
50 55 60

Glu Leu Val Ile Leu Thr Pro Pro Tyr Phe Ser Arg Ile Asn Ser Ile  
65 70 75 80

Asn Cys Ile Ser Cys Ser Arg Val Pro Tyr His Ser Leu Ser Leu Leu  
85 90 95

Thr Val Thr Ser Thr Pro Ala Ser Arg Asp Pro Met Leu Ala Thr Met  
100 105 110

Xaa

<210> 1772  
<211> 260  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> 6x FoxP3NFAT miniP

<400> 1772  
agaacatttc tctggcctaa ctggccggta ccggcttcat tttttccatt tactgcagag 60  
gcttcatttt ttccatttac tgcagaggct tcattttttc catttactgc agaggcttca 120  
ttttttccat ttactgcaga ggcttcattt tttccattta ctgcagaggc ttcatttttt 180  
ccatttactg cagaactagt agagggtata taatggaagc tcgacttcca gctcgagaga 240  
cccaatgcta gccaccatgg 260

<210> 1773  
<211> 86  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> 6x FoxP3NFAT miniP

<400> 1773

Arg Thr Phe Leu Trp Pro Asn Trp Pro Val Pro Ala Ser Phe Phe Pro  
1 5 10 15

Phe Thr Ala Glu Ala Ser Phe Phe Pro Phe Thr Ala Glu Ala Ser Phe  
20 25 30

Phe Pro Phe Thr Ala Glu Ala Ser Phe Phe Pro Phe Thr Ala Glu Ala  
35 40 45

Ser Phe Phe Pro Phe Thr Ala Glu Ala Ser Phe Phe Pro Phe Thr Ala  
50 55 60

Glu Leu Val Glu Gly Ile Trp Lys Leu Asp Phe Gln Leu Glu Arg Pro  
65 70 75 80

Asn Ala Ser His His Gly

<210> 1774  
 <211> 365  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> 6x IL2NFAT mCMV

<400> 1774  
 agaacatttc tctggcctaa ctggccggta ccggaggaaa aactgtttca tacagaaggc 60  
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 gtggaggaaa aactgtttca tacagaaggc gtggaggaaa aactgtttca tacagaaggc 180  
 gtggaggaaa aactgtttca tacagaaggc gtactagtta ggcgtgtacg gtgggaggcc 240  
 tatataagca gagctcgttt agtgaaccgt cagatcgctt ggagacgcca tccacgctgt 300  
 ttgacctcc atagaagaca ccgggaccga tccagcctcg agagacccaa tgctagccac 360  
 catgg 365

<210> 1775  
 <211> 122  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> 6x IL2NFAT mCMV

<400> 1775

Arg Thr Phe Leu Trp Pro Asn Trp Pro Val Pro Glu Glu Lys Leu Phe  
 1 5 10 15

His Thr Glu Gly Val Glu Glu Lys Leu Phe His Thr Glu Gly Val Glu  
 20 25 30

Glu Lys Leu Phe His Thr Glu Gly Val Glu Glu Lys Leu Phe His Thr  
 35 40 45

Glu Gly Val Glu Glu Lys Leu Phe His Thr Glu Gly Val Glu Glu Lys  
 50 55 60

Leu Phe His Thr Glu Gly Val Leu Val Arg Arg Val Arg Trp Glu Ala  
 65 70 75 80

Tyr Ile Ser Arg Ala Arg Leu Val Asn Arg Gln Ile Ala Trp Arg Arg  
 85 90 95

His Pro Arg Cys Phe Asp Leu His Arg Arg His Arg Asp Arg Ser Ser  
 100 105 110

Leu Glu Arg Pro Asn Ala Ser His His Gly  
 115 120

<210> 1776  
<211> 361  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> 6x IL2NFAT mIL2P

<400> 1776  
agaacatttc tctggcctaa ctggccggtc ccggaggaaa aactgtttca tacagaaggc 60  
gtggaggaaa aactgtttca tacagaaggc gtggaggaaa aactgtttca tacagaaggc 120  
gtggaggaaa aactgtttca tacagaaggc gtggaggaaa aactgtttca tacagaaggc 180  
gtggaggaaa aactgtttca tacagaaggc gtactagtca ttttgacacc cccataatat 240  
ttttccagaa ttaacagtat aaattgcatc tcttgttcaa gagttcccta tcactctctt 300  
taatcactac tcacagtaac ctcaactcct gcctcgagag acccaatgct agccaccatg 360  
g 361

<210> 1777  
<211> 119  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> 6x IL2NFAT mIL2P

<220>  
<221> misc\_feature  
<222> (119)..(119)  
<223> Xaa can be any naturally occurring amino acid

<400> 1777

Arg Thr Phe Leu Trp Pro Asn Trp Pro Val Pro Glu Glu Lys Leu Phe  
1 5 10 15

His Thr Glu Gly Val Glu Glu Lys Leu Phe His Thr Glu Gly Val Glu  
20 25 30

Glu Lys Leu Phe His Thr Glu Gly Val Glu Glu Lys Leu Phe His Thr  
35 40 45

Glu Gly Val Glu Glu Lys Leu Phe His Thr Glu Gly Val Glu Glu Lys  
50 55 60

Leu Phe His Thr Glu Gly Val Leu Val Ile Leu Thr Pro Pro Tyr Phe  
65 70 75 80

Ser Arg Ile Asn Ser Ile Asn Cys Ile Ser Cys Ser Arg Val Pro Tyr  
85 90 95

His Ser Leu Ser Leu Leu Thr Val Thr Ser Thr Pro Ala Ser Arg Asp  
100 105 110

Pro Met Leu Ala Thr Met Xaa

<210> 1778  
 <211> 278  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> 6x IL2NFAT miniP

<400> 1778  
 agaacatttc tctggcctaa ctggccggta ccggaggaaa aactgtttca tacagaaggc 60  
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 gtggaggaaa aactgtttca tacagaaggc gtggaggaaa aactgtttca tacagaaggc 180  
 gtggaggaaa aactgtttca tacagaaggc gtactagtag agggatatata atggaagctc 240  
 gacttccagc tcgagagacc caatgctagc caccatgg 278

<210> 1779  
 <211> 92  
 <212> PRT  
 <213> Artificial Sequence

<220>  
 <223> 6x IL2NFAT miniP

<400> 1779

Arg Thr Phe Leu Trp Pro Asn Trp Pro Val Pro Glu Glu Lys Leu Phe  
 1 5 10 15

His Thr Glu Gly Val Glu Glu Lys Leu Phe His Thr Glu Gly Val Glu  
 20 25 30

Glu Lys Leu Phe His Thr Glu Gly Val Glu Glu Lys Leu Phe His Thr  
 35 40 45

Glu Gly Val Glu Glu Lys Leu Phe His Thr Glu Gly Val Glu Glu Lys  
 50 55 60

Leu Phe His Thr Glu Gly Val Leu Val Glu Gly Ile Trp Lys Leu Asp  
 65 70 75 80

Phe Gln Leu Glu Arg Pro Asn Ala Ser His His Gly  
 85 90

<210> 1780  
 <211> 459  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Human NME1

<400> 1780  
 atggccaact gtgagcgtac cttcattgcg atcaaaccag atgggggtcca gcgggggtctt 60  
 gtgggagaga ttatcaagcg ttttgagcag aaaggattcc gccttggttg tctgaaattc 120



atgcaagctt ccgaagatct tctcaaggaa cactacgttg acctgaagga ccgtccattc 180  
tttgccggcc tggtgaaata catgcactca gggccggtag ttgccatggt ctgggagggg 240  
ctgaatgtgg tgaagacggg ccgagtcatg ctcggggaga ccaaccctgc agactccaag 300  
cctgggacca tccgtggaga cttctgcata caagttggca ggaacattat acatggcagt 360  
gattctgtgg agagtgcaga gaaggagatc ggcttgtggt ttcaccctga ggaactggta 420  
gattacacga gctgtgctca gaactggatc tatgaatga 459

<210> 1781  
<211> 152  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Human NME1

<400> 1781

Met Ala Asn Cys Glu Arg Thr Phe Ile Ala Ile Lys Pro Asp Gly Val  
1 5 10 15  
  
Gln Arg Gly Leu Val Gly Glu Ile Ile Lys Arg Phe Glu Gln Lys Gly  
20 25 30  
  
Phe Arg Leu Val Gly Leu Lys Phe Met Gln Ala Ser Glu Asp Leu Leu  
35 40 45  
  
Lys Glu His Tyr Val Asp Leu Lys Asp Arg Pro Phe Phe Ala Gly Leu  
50 55 60  
  
Val Lys Tyr Met His Ser Gly Pro Val Val Ala Met Val Trp Glu Gly  
65 70 75 80  
  
Leu Asn Val Val Lys Thr Gly Arg Val Met Leu Gly Glu Thr Asn Pro  
85 90 95  
  
Ala Asp Ser Lys Pro Gly Thr Ile Arg Gly Asp Phe Cys Ile Gln Val  
100 105 110  
  
Gly Arg Asn Ile Ile His Gly Ser Asp Ser Val Glu Ser Ala Glu Lys  
115 120 125  
  
Glu Ile Gly Leu Trp Phe His Pro Glu Glu Leu Val Asp Tyr Thr Ser  
130 135 140  
  
Cys Ala Gln Asn Trp Ile Tyr Glu  
145 150

<210> 1782  
<211> 1131  
<212> DNA  
<213> Artificial Sequence

<220>

<223> Human NME7

<400> 1782

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atgaatcata gtgaaagatt cgttttcatt gcagagtggg atgatccaaa tgcttcactt    60
cttcgacgtt atgagctttt attttaccca ggggatggat ctgttgaaat gcatgatgta    120
aagaatcatc gcaccttttt aaagcggacc aaatatgata acctgcactt ggaagattta    180
tttataggca acaaagtga tgtcttttct cgacaactgg tattaattga ctatggggat    240
caatatacag ctcgccagct gggcagtagg aaagaaaaaa cgctagccct aattaaacca    300
gatgcaatat caaaggctgg agaaataatt gaaataataa acaaagctgg atttactata    360
accaaactca aaatgatgat gctttcaagg aaagaagcat tggattttca tgtagatcac    420
cagtcaagac cttttttcaa tgagctgata cagtttatta caactgggtcc tattattgcc    480
atggagattt taagagatga tgctatatgt gaatggaaaa gactgctggg acctgcaaac    540
tctggagtgg cacgcacaga tgcttctgaa agcattagag ccctctttgg aacagatggc    600
ataagaaatg cagcgcatgg ccctgattct tttgcttctg cggccagaga aatggagttg    660
ttttttcctt caagtggagg ttgtgggccg gcaaacactg ctaaatttac taattgtacc    720
tgttgcattg ttaaacccca tgctgtcagt gaaggactgt tgggaaagat cctgatggct    780
atccgagatg caggttttga aatctcagct atgcagatgt tcaatatgga tcgggttaat    840
gttgaggaat tctatgaagt ttataaagga gtagtgaccg aatatcatga catggtgaca    900
gaaatgtatt ctggcccttg ttagcaatg gagattcaac agaataatgc taaaagaca    960
tttcgagaat tttgtggacc tgctgatact gaaattgccc ggcatattac ccctggaact   1020
ctcagagcaa tctttggtaa aactaagatc cagaatgctg ttcactgtac tgatctgcca   1080
gaggatggcc tattagaggt tcaatacttc ttcaagatct tggataatta g           1131
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<210> 1783

<211> 376

<212> PRT

<213> Artificial Sequence

<220>

<223> Human NME7

<400> 1783

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Met Asn His Ser Glu Arg Phe Val Phe Ile Ala Glu Trp Tyr Asp Pro
1           5           10          15
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Asn Ala Ser Leu Leu Arg Arg Tyr Glu Leu Leu Phe Tyr Pro Gly Asp
          20          25          30
```

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Gly Ser Val Glu Met His Asp Val Lys Asn His Arg Thr Phe Leu Lys
          35          40          45
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Arg Thr Lys Tyr Asp Asn Leu His Leu Glu Asp Leu Phe Ile Gly Asn
          50          55          60
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Lys Val Asn Val Phe Ser Arg Gln Leu Val Leu Ile Asp Tyr Gly Asp  
65 70 75 80

Gln Tyr Thr Ala Arg Gln Leu Gly Ser Arg Lys Glu Lys Thr Leu Ala  
85 90 95

Leu Ile Lys Pro Asp Ala Ile Ser Lys Ala Gly Glu Ile Ile Glu Ile  
100 105 110

Ile Asn Lys Ala Gly Phe Thr Ile Thr Lys Leu Lys Met Met Met Leu  
115 120 125

Ser Arg Lys Glu Ala Leu Asp Phe His Val Asp His Gln Ser Arg Pro  
130 135 140

Phe Phe Asn Glu Leu Ile Gln Phe Ile Thr Thr Gly Pro Ile Ile Ala  
145 150 155 160

Met Glu Ile Leu Arg Asp Asp Ala Ile Cys Glu Trp Lys Arg Leu Leu  
165 170 175

Gly Pro Ala Asn Ser Gly Val Ala Arg Thr Asp Ala Ser Glu Ser Ile  
180 185 190

Arg Ala Leu Phe Gly Thr Asp Gly Ile Arg Asn Ala Ala His Gly Pro  
195 200 205

Asp Ser Phe Ala Ser Ala Ala Arg Glu Met Glu Leu Phe Phe Pro Ser  
210 215 220

Ser Gly Gly Cys Gly Pro Ala Asn Thr Ala Lys Phe Thr Asn Cys Thr  
225 230 235 240

Cys Cys Ile Val Lys Pro His Ala Val Ser Glu Gly Leu Leu Gly Lys  
245 250 255

Ile Leu Met Ala Ile Arg Asp Ala Gly Phe Glu Ile Ser Ala Met Gln  
260 265 270

Met Phe Asn Met Asp Arg Val Asn Val Glu Glu Phe Tyr Glu Val Tyr  
275 280 285

Lys Gly Val Val Thr Glu Tyr His Asp Met Val Thr Glu Met Tyr Ser  
290 295 300

Gly Pro Cys Val Ala Met Glu Ile Gln Gln Asn Asn Ala Thr Lys Thr  
305 310 315 320

Phe Arg Glu Phe Cys Gly Pro Ala Asp Pro Glu Ile Ala Arg His Leu  
325 330 335

Arg Pro Gly Thr Leu Arg Ala Ile Phe Gly Lys Thr Lys Ile Gln Asn

Ala Val His Cys Thr Asp Leu Pro Glu Asp Gly Leu Leu Glu Val Gln  
 355 360 365

Tyr Phe Phe Lys Ile Leu Asp Asn  
 370 375

<210> 1784  
 <211> 1359  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Humanized C2 IgG1 heavy chain sequence

<400> 1784  
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 ccaggaagg ggctggagt ggtctcaacc attagtagtg gcggaaccta catatactac 180  
 cccgactcag tgaagggccg attcaccatc tccagagaca acgccaagaa ctactgtat 240  
 ctgcaaatga acagcctgag agccgaggac acggccgtgt attactgtgc gagacttggg 300  
 ggggataatt actacgaata cttcgatgtc tggggcaaaag ggaccacggc caccgtctcc 360  
 tccgctagca ccaagggcc atcgggtcttc cccctggcac cctcctcaa gagcacctct 420  
 gggggcacag cggccctggg ctgcctggc aaggactact tccccgaacc ggtgacgggtg 480  
 tcgtggaact caggcgccct gaccagcggc gtgcacacct tcccggctgt cctacagtcc 540  
 tcaggactct actccctcag cagcgtgggt acagtgcct ccagcagctt gggcacccag 600  
 acctacatct gcaacgtgaa tcacaagccc agcaacacca aggtggacaa gaaagttgag 660  
 cccaaatctt gtgacaaaac tcacacatgc ccaccgtgcc cagcacctga actcctgggg 720  
 ggaccgtcag tcttctctt cccccaaaa cccaaggaca ccctcatgat ctcccggacc 780  
 cctgaggtca catgcgtggg ggtggacgtg agccacgaag accctgaggt caagttcaac 840  
 tgggtacgtgg acggcgtgga ggtgcataat gccaaagaaa agccgcggga ggagcagtac 900  
 aacagcacgt accgtgtggg cagcgtcctc accgtcctgc accaggactg gctgaatggc 960  
 aaggagtaca agtgaagggt ctccaacaaa gccctcccag ccccatcga gaaaaccatc 1020  
 tccaaagcca aagggcagcc ccgagaacca caggtgtaca ccctgcccc atcccgggag 1080  
 gagatgacca agaaccagggt cagcctgacc tgcctgggtc aaggcttcta tcccagcgac 1140  
 atcggcgtgg agtgggagag caatgggcag ccggagaaca actacaagac cagcctccc 1200  
 gtgctggact ccgacggctc cttcttcctc tacagcaagc tcaccgtgga caagagcagg 1260  
 tggcagcagg ggaacgtctt ctcatgtctc gtgatgcatg aggctctgca caaccactac 1320  
 acgcagaaga gcctctccct gtctccgggt aatgataa 1359

<210> 1785  
 <211> 12

<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 17H6 heavy chain variable complementarity determining regions 3 (CDR3) sequence

<400> 1785

Asp Tyr Tyr Gly Ser Asn Pro Ala Trp Phe Ala Tyr  
1 5 10

<210> 1786  
<211> 11  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Mouse 17H6 heavy chain variable framework 4 (FW4) sequence

<400> 1786

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ala  
1 5 10

<210> 1787  
<211> 72  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> CD8 transmembrane domain

<400> 1787  
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accctttact gc 72

<210> 1788  
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