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(54) Title: COMBINATION OF EPIGENETIC FACTORS AND BISPECIFIC COMPOUNDS TARGETING CD33 AND CD3 IN THE TREATMENT OF MYELOID LEUKEMIA

(57) Abstract: The present invention provides a combination epigenetic factors and bispecific compounds targeting CD33 and CD3 in the treatment of myeloid leukemia, wherein the epigenetic factor is selected from the group consisting of histone deacetylase (HDAC) inhibitors, DNA methyltransferase (DNMT) I inhibitors, hydroxyurea, Granulocyte-Colony Stimulating Factor (G-CSF), histone demethylase inhibitors and ATRA (All Trans-retinoic acid). Accordingly, the invention provides a pharmaceutical composition comprising a CD33 targeting compound and at least one epigenetic factor and an epigenetic factor for use in the amelioration and/or treatment of a myeloid leukemia, wherein the epigenetic factor increases the responsiveness of a patient to a CD33 targeting compound. Moreover, the invention provides the use of at least one an epigenetic factor for increasing the responsiveness of a myeloid leukemia patient to a treatment with a CD33 targeting compound, a method for the treatment of a myeloid leukemia, the method comprising the administration of at least one epigenetic factor and a CD33 targeting compound to a patient in the need thereof and a kit comprising a pharmaceutical composition of the invention or an epigenetic factor of the invention and a bispecific CD33 targeting compound.



**Combination of epigenetic factors and bispecific compounds targeting CD33 and CD3
in the treatment of myeloid leukemia**

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Field of the Invention

The present invention provides a combination epigenetic factors and bispecific compounds targeting CD33 and CD3 in the treatment of myeloid leukemia, wherein the epigenetic factor is selected from the group consisting of histone deacetylase (HDAC) inhibitors, DNA methyltransferase (DNMT) I inhibitors, hydroxyurea, Granulocyte-Colony Stimulating Factor (G-CSF), histone demethylase inhibitors and ATRA (All Trans-retinoic acid). Accordingly, the invention provides a pharmaceutical composition comprising a CD33 targeting compound and at least one epigenetic factor and an epigenetic factor for use in the amelioration, and/or treatment of a myeloid leukemia, wherein the epigenetic factor increases the responsiveness of a patient to a CD33 targeting compound. Moreover, the invention provides the use of at least one an epigenetic factor for increasing the responsiveness of a myeloid leukemia patient to a treatment with a CD33 targeting compound, a method for the treatment of a myeloid leukemia, the method comprising the administration of at least one epigenetic factor and a CD33 targeting compound to a patient in the need thereof and a kit comprising a pharmaceutical composition of the invention or an epigenetic factor of the invention and a bispecific CD33 targeting compound.

Background of the invention

Despite some gradual improvements over the last decades with the use of intensive therapies – including multiagent conventional chemotherapy and allogeneic stem cell transplantation (SCT) – the survival of patients with acute myeloid leukemia (AML) remains poor, and patient are at high risk of experiencing treatment-related morbidity and mortality. For example, although SCT has proven to provide a potent anti-leukemic effect that can lead to elimination of chemoresistant leukemic cells, a large number of patients will develop significant graft versus host (GvH) disease that will eventually be fatal in many. For many patients, in particular those who are not suitable for such intensive therapeutic strategies, novel therapeutic options including immunotherapeutic approaches are urgently sought after. A promising immunotherapeutic strategy devoid of GvH reactions is to recruit in-vivo the patient's own T-cells and retarget them directly at leukemic cells.

This approach became feasible by a novel class of bispecific single-chain antibodies, which direct cytotoxic T-lymphocytes at predefined surface antigens on tumor cells (Baeuerle et al. Curr Opin Mol Ther. 2009;11:22-30). Clinical proof of concept was provided by

blinatumomab, a bispecific antibody directed at both the CD19 B-cell surface antigen and the CD3 ϵ component of the T-cell receptor complex. Its therapeutic efficacy was shown for patients with B-cell lymphomas and B precursor acute lymphoblastic leukemia (ALL).

Acute myeloid leukemia (AML) has served as paradigm for the therapeutic use of monoclonal antibodies because of well-defined cell surface antigens and easy tumor accessibility. The most investigated target so far is CD33, a sialic-acid-dependent cytoadhesion molecule known as a myeloid differentiation antigen found on AML blasts in most patients and, perhaps, leukemic stem cells in some. While the intact humanized CD33 antibody lintuzumab induced complete remission in individual patients in a single-agent phase I and II trial, no survival benefit was found in a phase III trial when it was combined with a triple chemotherapy regimen.

Summary of the invention

The present invention provides a pharmaceutical composition comprising a CD33 targeting compound and at least one epigenetic factor, wherein

- (a) the CD33 targeting compound is a bispecific construct comprising a first binding domain specifically binding to CD33 and a second binding domain specifically binding to CD3; and
- (b) the at least one epigenetic factor is selected from the group consisting of histone deacetylase (HDAC) inhibitors, DNA methyltransferase (DNMT) I inhibitors, hydroxyurea, Granulocyte-Colony Stimulating Factor (G-CSF), histone methyltransferase (HMT) inhibitors and ATRA (All Trans-retinoic acid).

In one embodiment the pharmaceutical composition of the invention is characterized in a way that the at least one epigenetic factor is selected from the group comprising:

- (a) a histone deacetylase (HDAC) inhibitor selected from the group consisting of panobinostat, vorinostat, romidepsin, N-acetyldinaline, belinostat, givinostat, entinostat, mocetinostat, EVP-0334, SRT501, CUDC-101, Quisinostat, abexinostat, LAQ824, and valproic acid;
- (b) a DNA methyltransferase (DNMT) I inhibitor selected from the group consisting of 5-azacitidine, decitabine, hydralazine, zebularine, procainamide, (-)-epigallocatechin-3-gallate, MG98, RG108, and SGI-110; and
- (c) a histone methyltransferase (HMT) inhibitor selected from the group consisting of LSD1 (KDM1A) demethylase inhibitor, and chaetocin.

The invention also provides a pharmaceutical composition, wherein

- (a) the epigenetic factor is administered prior to the administration of the CD33 targeting compound;

- (b) the epigenetic factor is administered subsequent to the administration of the CD33 targeting compound; or
- (c) the epigenetic factor and the CD33 targeting compound are administered simultaneously.

- 5 The invention further provides a pharmaceutical composition comprising a CD33 targeting compound and at least one epigenetic factor, wherein (a) the CD33 targeting compound is a bispecific single chain antibody construct comprising a first binding domain specifically binding to CD33 and a second binding domain specifically binding to CD3; and (b) the at least one epigenetic factor is selected from
- 10 the group consisting of histone deacetylase (HDAC) inhibitor panobinostat, DNA methyltransferase (DNMT) I inhibitor 5-azacitidine, hydroxyurea, and Granulocyte-Colony Stimulating Factor (G-CSF), and wherein (i) the epigenetic factor is administered prior to the administration of the CD33 targeting compound; (ii) the epigenetic factor is administered subsequent to the administration of the CD33
- 15 targeting compound; or (iii) the epigenetic factor and the CD33 targeting compound are administered simultaneously.

The invention further provides a pharmaceutical composition, wherein a first dose of the epigenetic factor is administered prior to the start of the administration of the CD33 targeting compound.

- 20 Moreover, the invention further provides a pharmaceutical composition, wherein the administration of the epigenetic factor is continued during the administration of the CD33 targeting compound.

In one embodiment the pharmaceutical composition of the invention is characterized in a way that the bispecific construct is a bispecific antibody construct.

- 25 In preferred embodiment of the pharmaceutical composition of the invention the bispecific antibody construct is a bispecific single chain antibody construct.

It is also preferred for the pharmaceutical composition of the invention that the bispecific antibody construct binds to human and cynomolgous CD3 and human and cynomolgous CD33.

- 30 In one embodiment of the pharmaceutical composition of invention the bispecific antibody construct comprises

- a first binding domain specifically binding to CD33 and comprising a VL chain having an amino acid sequence as depicted in SEQ ID NO: 6, 24, 42, 60, 78, 96, 114 and 132 and a VH chain having an amino acid sequence as depicted in SEQ
- 35 ID NO: 1, 19, 37, 55, 73, 91, 109 and 127; and
- a second binding domain specifically binding to CD3 and comprising a VL chain having an amino acid sequence as depicted in SEQ ID NO: 154, 157, 160, 163,

166, 169 and 172 and a VH chain having an amino acid sequence as depicted in SEQ ID NO:155, 158, 161, 164, 167, 170 and 173.

In preferred embodiment of the pharmaceutical composition the bispecific antibody construct comprises an amino acid sequence as depicted in any of SEQ ID NOs: 13, 15, 17, 31, 33, 35, 49, 51, 53, 67, 69, 71, 85, 87, 89, 103, 105, 107, 121, 123, 125, 139, 141, 143, 148, 150, 152, 215, 217, 219, 221, 223, 225 and 227.

The pharmaceutical composition of the invention is envisaged for the treatment of CD33 expressing myeloid leukemia. Preferably, the myeloid leukemia is selected from the group consisting of acute myeloblastic leukemia, chronic neutrophilic leukemia, myeloid dendritic cell leukemia, accelerated phase chronic myelogenous leukemia, acute myelomonocytic leukemia, juvenile myelomonocytic leukemia, chronic myelomonocytic leukemia, acute basophilic leukemia, acute eosinophilic leukemia, chronic eosinophilic leukemia, Acute

[Text continued on page 4]

megakaryoblastic leukemia, essential thrombocytosis, acute erythroid leukemia, polycythemia vera, myelodysplastic syndrome, acute panmyelosis, myeloid sarcoma, and acute biphenotypic leukaemia.

In an alternative embodiment the invention provides an epigenetic factor for use in the amelioration and/or treatment of a myeloid leukemia, wherein the epigenetic factor increases the responsiveness of a patient to a CD33 targeting compound, wherein

(a) the CD33 targeting compound is a bispecific construct comprising a first binding domain specifically binding to CD33 and a second binding domain specifically binding to CD3; and

(b) the at least one epigenetic factor is selected from the group consisting of histone deacetylase (HDAC) inhibitors, DNA methyltransferase (DNMT) I inhibitors, hydroxyurea, Granulocyte-Colony Stimulating Factor (G-CSF), histone methyltransferase (HMT) inhibitors and ATRA (All Trans-retinoic acid).

In one embodiment of the epigenetic factor of the invention the epigenetic factor is selected from the group comprising

(a) a histone deacetylase (HDAC) inhibitor selected from the group consisting of panobinostat, vorinostat, romidepsin, N-acetyldinaline, belinostat, givinostat, entinostat, mocetinostat, EVP-0334, SRT501, CUDC-101, Quisinostat, abexinostat, LAQ824, and valproic acid;

(b) a DNA methyltransferase (DNMT) I inhibitor selected from the group consisting of 5-azacitidine, decitabine, hydralazine, zebularine, procainamide, (-)-epigallocatechin-3-gallate, MG98, RG108, and SGI-110; and

(c) a histone methyltransferase (HMT) inhibitor selected from the group consisting of LSD1 (KDM1A) demethylase inhibitor, and chaetocin.

Moreover, in one embodiment of the epigenetic factor of the invention

(a) the epigenetic factor is administered prior to the administration of the CD33 targeting compound;

(b) the epigenetic factor is administered subsequent to the administration of the CD33 targeting compound; or

(c) the epigenetic factor and the CD33 targeting compound are administered simultaneously.

It is preferred for the epigenetic factor of the invention that a first dose of the epigenetic factor is administered prior to the start of the administration of the CD33 targeting compound.

Preferably, the administration of the epigenetic factor of the invention is continued during the administration of the CD33 targeting compound.

It is also preferred for the epigenetic factor the invention that the bispecific construct is a bispecific antibody construct.

In one embodiment of the epigenetic factor of the invention the bispecific antibody construct is a bispecific single chain antibody construct.

- 5 It is preferred for the epigenetic factor of the invention that the bispecific antibody construct binds to human and cynomolgous CD3 and CD33.

Also preferred for the epigenetic factor of the invention is that the bispecific antibody construct comprises

- a first binding domain specifically binding to CD33 and comprising a VL chain having an amino acid sequence as depicted in SEQ ID NO: 6, 24, 42, 60, 78, 96, 114 and 132 and a VH chain having an amino acid sequence as depicted in SEQ ID NO: 1, 19, 37, 55, 73, 91, 109 and 127; and
- a second binding domain specifically binding to CD3 and comprising a VL chain having an amino acid sequence as depicted in SEQ ID NO: 154, 157, 160, 163, 166, 169 and 172 and a VH chain having an amino acid sequence as depicted in SEQ ID NO: 155, 158, 161, 164, 167, 170 and 173.

In one embodiment of the epigenetic factor of the invention the bispecific antibody construct comprises an amino acid sequence as depicted in any of SEQ ID NOs: 13, 15, 17, 31, 33, 35, 49, 51, 53, 67, 69, 71, 85, 87, 89, 103, 105, 107, 121, 123, 125, 139, 141, 143, 148, 150, 152, 215, 217, 219, 221, 223, 225 and 227.

In one embodiment of the epigenetic factor of the invention the myeloid leukemia is selected from the group consisting of acute myeloblastic leukemia, chronic neutrophilic leukemia, myeloid dendritic cell leukemia, accelerated phase chronic myelogenous leukemia, acute myelomonocytic leukemia, juvenile myelomonocytic leukemia, chronic myelomonocytic leukemia, acute basophilic leukemia, acute eosinophilic leukemia, chronic eosinophilic leukemia, Acute megakaryoblastic leukemia, essential thrombocytosis, acute erythroid leukemia, polycythemia vera, myelodysplastic syndrome, acute panmyelosis, myeloid sarcoma, and acute biphenotypic leukaemia.

An alternative embodiment of the invention provides a use of at least one an epigenetic factor for increasing the responsiveness of a myeloid leukemia patient to a treatment with a CD33 targeting compound, wherein

- (a) the CD33 targeting compound is a bispecific construct comprising a first binding domain specifically binding to CD33 and a second binding domain specifically binding to CD3; and
- (b) the at least one epigenetic factor is selected from the group consisting of histone deacetylase (HDAC) inhibitors, DNA methyltransferase (DNMT) I inhibitors,

hydroxyurea, Granulocyte-Colony Stimulating Factor (G-CSF), histone demethylase inhibitors and ATRA (All Trans-retinoic acid).

In one embodiment of the use of the invention the at least one epigenetic factor is selected from the group comprising

- 5 (a) a histone deacetylase (HDAC) inhibitor selected from the group consisting of panobinostat, vorinostat, romidepsin, N-acetyldinaline, belinostat, givinostat, entinostat, mocetinostat, EVP-0334, SRT501, CUDC-101, Quisinostat, abexinostat, LAQ824, and valproic acid;
- 10 (b) a DNA methyltransferase (DNMT) I inhibitor selected from the group consisting of 5-azacitidine, decitabine, hydralazine, zebularine, procainamide, (-)-epigallocatechin-3-gallate, MG98, RG108, and SGI-110; and
- (c) a histone methyltransferase (HMT) inhibitor selected from the group consisting of LSD1 (KDM1A) demethylase inhibitor, and chaetocin.

Also in an embodiment of the use of the invention

- 15 (a) the epigenetic factor is administered prior to the administration of the CD33 targeting compound;
- (b) the epigenetic factor is administered subsequent to the administration of the CD33 targeting compound; or
- 20 (c) the epigenetic factor and the CD33 targeting compound are administered simultaneously.

In a preferred use of any of the invention a first dose of the epigenetic factor is administered prior to the start of the administration of the CD33 targeting compound.

Moreover, in one embodiment of the use of the invention the administration of the epigenetic factor is continued during the administration of the CD33 targeting compound.

- 25 In one embodiment of the use of the invention the bispecific construct is a bispecific antibody construct.

In a preferred embodiment of the use of the invention the bispecific antibody construct is a bispecific single chain antibody construct.

- 30 In one embodiment of the use of the invention the bispecific antibody construct binds to human and cynomolgous CD3 and CD33.

According to one embodiment of the use of the invention the bispecific antibody construct comprises

- a first binding domain specifically binding to CD33 and comprising a VL chain having an amino acid sequence as depicted in SEQ ID NO: 6, 24, 42, 60, 78, 96, 114 and 132 and a
- 35 VH chain having an amino acid sequence as depicted in SEQ ID NO: 1, 19, 37, 55, 73, 91, 109 and 127; and

- a second binding domain specifically binding to CD3 and comprising a VL chain having an amino acid sequence as depicted in SEQ ID NO: 154, 157, 160, 163, 166, 169 and 172 and a VH chain having an amino acid sequence as depicted in SEQ ID NO: 155, 158, 161, 164, 167, 170 and 173.

5 In a preferred embodiment of the use of the invention the bispecific antibody construct comprises an amino acid sequence as depicted in any of SEQ ID NOs: 13, 15, 17, 31, 33, 35, 49, 51, 53, 67, 69, 71, 85, 87, 89, 103, 105, 107, 121, 123, 125, 139, 141, 143, 148, 150, 152, 215, 217, 219, 221, 223, 225 and 227.

Furthermore, in an embodiment of the use of the invention the myeloid leukemia is selected
 10 from the group consisting of acute myeloblastic leukemia, chronic neutrophilic leukemia, myeloid dendritic cell leukemia, accelerated phase chronic myelogenous leukemia, acute myelomonocytic leukemia, juvenile myelomonocytic leukemia, chronic myelomonocytic leukemia, acute basophilic leukemia, acute eosinophilic leukemia, chronic eosinophilic leukemia, Acute megakaryoblastic leukemia, essential thrombocytosis, acute erythroid
 15 leukemia, polycythemia vera, myelodysplastic syndrome, acute panmyelosis, myeloid sarcoma, and acute biphenotypic leukaemia.

In a further alternative embodiment the invention provides a method for the treatment of a myeloid leukemia, the method comprising the administration of at least one epigenetic factor and a CD33 targeting compound to a patient in the need thereof, wherein

- 20 (a) the CD33 targeting compound is a bispecific construct comprising a first binding domain specifically binding to CD33 and a second binding domain specifically binding to CD3; and
- (b) the at least one epigenetic factor is selected from the group consisting of histone deacetylase (HDAC) inhibitors, DNA methyltransferase (DNMT) I inhibitors,
 25 hydroxyurea, Granulocyte-Colony Stimulating Factor (G-CSF), histone methyltransferase (HMT) inhibitors and ATRA (All Trans-retinoic acid).

In an embodiment of the method of the invention the at least one epigenetic factor is selected from the group comprising

- 30 (a) a histone deacetylase (HDAC) inhibitor selected from the group consisting of panobinostat, vorinostat, romidepsin, N-acetyldinaline, belinostat, givinostat, entinostat, mocetinostat, EVP-0334, SRT501, CUDC-101, Quisinostat, abexinostat, LAQ824, and valproic acid;
- (b) a DNA methyltransferase (DNMT) I inhibitor selected from the group consisting of 5-azacitidine, decitabine, hydralazine, zebularine, procainamide, (-)-epigallocatechin-3-
 35 gallate, MG98, RG108, and SGI-110; and
- (c) a histone methyltransferase (HMT) inhibitor selected from the group consisting of LSD1 (KDM1A) demethylase inhibitor, and chaetocin.

Also in an embodiment of the method of the invention

- (a) the epigenetic factor is administered prior to the administration of the CD33 targeting compound;
- (b) the epigenetic factor is administered subsequent to the administration of the CD33 targeting compound; or
- (c) the epigenetic factor and the CD33 targeting compound are administered simultaneously.

In a preferred embodiment of the method of the invention a first dose of the epigenetic factor is administered prior to the start of the administration of the CD33 targeting compound.

It is also preferred for the method of the invention that the administration of the epigenetic factor is continued during the administration of the CD33 targeting compound.

In one embodiment of the method of the invention the bispecific construct is a bispecific antibody construct.

It is preferred for the method of the invention that the bispecific antibody construct is a bispecific single chain antibody construct.

It is more preferred for the method of the invention that the bispecific antibody construct binds to human and cynomolgous CD3 and CD33.

In one embodiment of the method of the invention the bispecific antibody construct comprises

- a first binding domain specifically binding to CD33 and comprising a VL chain having an amino acid sequence as depicted in SEQ ID NO: 6, 24, 42, 60, 78, 96, 114 and 132 and a VH chain having an amino acid sequence as depicted in SEQ ID NO: 1, 19, 37, 55, 73, 91, 109 and 127; and
- a second binding domain specifically binding to CD3 and comprising a VL chain having an amino acid sequence as depicted in SEQ ID NO: 154, 157, 160, 163, 166, 169 and 172 and a VH chain having an amino acid sequence as depicted in SEQ ID NO: 155, 158, 161, 164, 167, 170 and 173.

Also in one embodiment of the method of the invention the bispecific antibody construct comprises an amino acid sequence as depicted in any of SEQ ID NOs: 13, 15, 17, 31, 33, 35, 49, 51, 53, 67, 69, 71, 85, 87, 89, 103, 105, 107, 121, 123, 125, 139, 141, 143, 148, 150, 152, 215, 217, 219, 221, 223, 225 and 227.

Furthermore, in one embodiment of the method of the invention the myeloid leukemia is selected from the group consisting of acute myeloblastic leukemia, chronic neutrophilic leukemia, myeloid dendritic cell leukemia, accelerated phase chronic myelogenous leukemia, acute myelomonocytic leukemia, juvenile myelomonocytic leukemia, chronic myelomonocytic leukemia, acute basophilic leukemia, acute eosinophilic leukemia, chronic eosinophilic leukemia, Acute megakaryoblastic leukemia, essential thrombocytosis, acute

erythroid leukemia, polycythemia vera, myelodysplastic syndrome, acute panmyelosis, myeloid sarcoma, and acute biphenotypic leukaemia.

A further alternative embodiment of the invention provides a kit comprising a pharmaceutical composition of the invention or an epigenetic factor of the invention and a bispecific CD33 targeting compound comprising a first binding domain specifically binding to CD33 and a second binding domain specifically binding to CD3, to be employed in the amelioration and/or treatment of a myeloid leukemia.

Brief description of the drawings

Figure 1:

Effect of panobinostat pretreatment on CD33 expression and AMG 330-induced cytotoxicity. Parental OCI-AML3 and KG-1a cells were either left untreated or pretreated with panobinostat for 72 hours. Subsequently, CD33 expression was quantified, and cells treated with/without AMG 330 (0-250 pg/mL) and various effector:target (E:T) cell ratios using healthy donor T-cells. 48 hours later, cell counts were determined and cytotoxicity was assessed with DAPI staining to quantify drug-specific cytotoxicity. Results are shown as mean $\pm$ SEM from 3 independent experiments performed in duplicate wells using a single healthy donor as source for exogenous T-cells.

Figure 2:

Effect of azacitidine pretreatment on CD33 expression and AMG 330-induced cytotoxicity. Parental KG-1a cells were either left untreated or pretreated with azacitidine for 72 hours. Subsequently, CD33 expression was quantified, and cells treated with/without AMG 330 (0-250 pg/mL) and various effector:target (E:T) cell ratios using healthy donor T-cells. 48 hours later, cell counts were determined and cytotoxicity was assessed with DAPI staining to quantify drug-specific cytotoxicity. Results are shown as mean $\pm$ SEM from 3 independent experiments performed in duplicate wells using a single healthy donor as source for exogenous T-cells.

Figure 3:

Hydroxyurea-concentration dependant up-regulation of CD33 on A) HL-60 and B) PL21 AML cells.

Figure 4:

Hydroxyurea dependant up-regulation of CD33 on primary AML cells from patient 2.

Figure 5:

G-CSF dependant up-regulation of CD33 on KG1 α AML cells after 10 days of incubation.

Figure 6:

G-CSF dependant up-regulation of CD33 on primary AML cells from patient 1 after 10 days of incubation.

Detailed description of the invention

Definitions:

It must be noted that as used herein, the singular forms "a", "an", and "the", include plural references unless the context clearly indicates otherwise. Thus, for example, reference to "a reagent" includes one or more of such different reagents and reference to "the method" includes reference to equivalent steps and methods known to those of ordinary skill in the art that could be modified or substituted for the methods described herein.

Unless otherwise indicated, the term "at least" preceding a series of elements is to be understood to refer to every element in the series. 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 described herein. Such equivalents are intended to be encompassed by the present invention.

The term "and/or" wherever used herein includes the meaning of "and", "or" and "all or any other combination of the elements connected by said term".

The term "about" or "approximately" as used herein means within $\pm 20\%$, preferably within $\pm 15\%$, more preferably within $\pm 10\%$, and most preferably within $\pm 5\%$ of a given value or range.

Throughout this specification and the claims which follow, unless the context requires otherwise, the word "comprise", and variations such as "comprises" and "comprising", will be understood to imply the inclusion of a stated integer or step or group of integers or steps but not the exclusion of any other integer or step or group of integer or step. When used herein the term "comprising" can be substituted with the term "containing" or "including" or sometimes when used herein with the term "having".

When used herein "consisting of" excludes any element, step, or ingredient not specified in the claim element. When used herein, "consisting essentially of" does not exclude materials or steps that do not materially affect the basic and novel characteristics of the claim.

In each instance herein any of the terms "comprising", "consisting essentially of" and "consisting of" may be replaced with either of the other two terms.

The term "CD33 targeting compound" defines in the context of the present invention a binding molecule capable of specifically binding to the extracellular domain of the cell surface molecule CD33. As mentioned above CD33, a sialic-acid-dependent cytoadhesion molecule, is known as a myeloid differentiation antigen found on AML blasts in most patients. As apparent from the description of the embodiments of the invention, the CD33

targeting compound in the sense of the invention is a bispecific compound comprising at least a binding domain for the extracellular domain of the cell surface molecule CD33 and the extracellular domain of the molecule CD3 expressed on T cells. It is emphasized that the CD33 targeting compound comprises in the sense of the invention at least the binding domain for CD33 and CD3 and may also comprise one or more additional binding domains for other target structures, which result in a tri- or multispecific compound. In other words, the term "CD33 targeting compound" in the sense of the present disclosure indicates any molecule capable of (specifically) binding to, interacting with or recognizing the target molecule CD33 on the surface of a target cell and CD3 on the surface of a T cell. Such molecules or constructs may include proteinaceous parts and non-proteinaceous parts (e.g. chemical linkers or chemical cross-linking agents such as glutaraldehyde).

The term "binding molecule" in the sense of the present disclosure indicates any molecule capable of (specifically) binding to, interacting with or recognizing the target molecule.

A binding molecule, so to say, provides the scaffold for said one or more binding domains so that said binding domains can bind/interact with the surface molecule on a target cell and CD3 receptor complex on a T cell. For example, such a scaffold could be provided by protein A, in particular, the Z-domain thereof (affibodies), ImmE7 (immunity proteins), BPTI/APPI (Kunitz domains), Ras-binding protein AF-6 (PDZ-domains), charybdotoxin (Scorpion toxin), CTLA-4, Min-23 (knottins), lipocalins (anticalins), neokarzinostatin, a fibronectin domain, an ankyrin consensus repeat domain (Stumpp et al., Curr Opin Drug Discov Devel. 10(2), 153-159 (2007)) or thioredoxin (Skerra, Curr. Opin. Biotechnol. 18, 295-304 (2005); Hosse et al., Protein Sci. 15, 14-27 (2006); Nicaise et al., Protein Sci. 13, 1882-1891 (2004); Nygren and Uhlen, Curr. Opin. Struc. Biol. 7, 463-469 (1997)). A preferred binding molecule is an antibody, more preferably a bispecific antibody.

The term "single chain binding molecule" defines in connection with the present invention that the disclosed binding molecules in its simplest form are monomers. The molecules or constructs may include proteinaceous parts and non-proteinaceous parts (e.g. chemical linkers or chemical cross-linking agents such as glutaraldehyde). Thus, the single chain binding molecule may comprising accordance with the invention non-peptide linkers preferably to link at least two of the binding domains. Also in line with this invention are herein defined peptide linkers.

The definition of the term "antibody" includes embodiments such as monoclonal, chimeric, single chain, humanized and human antibodies, as well as antibody fragments, like, inter alia, Fab fragments. Antibody fragments or derivatives further comprise F(ab')₂, Fv, scFv fragments or single domain antibodies such as domain antibodies or nanobodies, single

variable domain antibodies or immunoglobulin single variable domain comprising merely one variable domain, which might be VHH, VH or VL, that specifically bind an antigen or epitope independently of other V regions or domains; see, for example, Harlow and Lane (1988) and (1999), loc. cit.; Kontermann and Dübel, Antibody Engineering, Springer, 2nd ed. 2010 and
5 Little, Recombinant Antibodies for Immunotherapy, Cambridge University Press 2009. Such immunoglobulin single variable domain encompasses not only an isolated antibody single variable domain polypeptide, but also larger polypeptides that comprise one or more monomers of an antibody single variable domain polypeptide sequence.

Monovalent antibody fragments in line with the above definition describe an embodiment of a
10 binding domain in connection with this invention. Such monovalent antibody fragments bind to a specific antigen and can be also designated "antigen-binding domain", "antigen-binding fragment" or "antibody binding region".

In line with this definition all above described embodiments of the term antibody can be subsumed under the term "antibody construct". Said term also includes diabodies or Dual-
15 Affinity Re-Targeting (DART) antibodies. Further envisaged are (bispecific) single chain diabodies, tandem diabodies (Tandab's), „minibodies“ exemplified by a structure which is as follows: (VH-VL-CH3)₂, (scFv-CH3)₂ or (scFv-CH3-scFv)₂, „Fc DART“ antibodies and „IgG DART“ antibodies, and multibodies such as triabodies. Immunoglobulin single variable domains encompass not only an isolated antibody single variable domain polypeptide, but
20 also larger polypeptides that comprise one or more monomers of an antibody single variable domain polypeptide sequence.

Various procedures are known in the art and may be used for the production of such antibody constructs (antibodies and/or fragments). Thus, (antibody) derivatives can be produced by peptidomimetics. Further, techniques described for the production of single
25 chain antibodies (see, inter alia, US Patent 4,946,778, Kontermann and Dübel (2010), loc. cit. and Little(2009), loc. cit.) can be adapted to produce single chain antibodies specific for elected polypeptide(s). Also, transgenic animals may be used to express humanized antibodies specific for polypeptides and fusion proteins of this invention. For the preparation of monoclonal antibodies, any technique, providing antibodies produced by continuous cell
30 line cultures can be used. Examples for such techniques include the hybridoma technique (Köhler and Milstein Nature 256 (1975), 495-497), the trioma technique, the human B-cell hybridoma technique (Kozbor, Immunology Today 4 (1983), 72) and the EBV-hybridoma technique to produce human monoclonal antibodies (Cole et al., Monoclonal Antibodies and Cancer Therapy, Alan R. Liss, Inc. (1985), 77-96). Surface plasmon resonance as employed
35 in the BIAcore system can be used to increase the efficiency of phage antibodies which bind to an epitope of a target polypeptide, such as CD3 epsilon (Schier, Human Antibodies

Hybridomas 7 (1996), 97-105; Malmborg, J. Immunol. Methods 183 (1995), 7-13). It is also envisaged in the context of this invention that the term "antibody" comprises antibody constructs, which may be expressed in a host as described herein below, e.g. antibody constructs which may be transfected and/or transduced via, inter alia, viruses or plasmid vectors.

Furthermore, the term "antibody" as employed herein also relates to derivatives or variants of the antibodies described herein which display the same specificity as the described antibodies. Examples of "antibody variants" include humanized variants of non-human antibodies, "affinity matured" antibodies (see, e.g. Hawkins et al. J. Mol. Biol. 254, 889-896 (1992) and Lowman et al., Biochemistry 30, 10832- 10837 (1991)) and antibody mutants with altered effector function(s) (see, e.g., US Patent 5, 648, 260, Kontermann and Dübel (2010), loc. cit. and Little(2009), loc. cit.).

The terms "antigen-binding domain", "antigen-binding fragment" and "antibody binding region" when used herein refer to a part of an antibody molecule that comprises amino acids responsible for the specific binding between antibody and antigen. The part of the antigen that is specifically recognized and bound by the antibody is referred to as the "epitope" as described herein above. As mentioned above, an antigen-binding domain may typically comprise an antibody light chain variable region (VL) and an antibody heavy chain variable region (VH); however, it does not have to comprise both. Fd fragments, for example, have two VH regions and often retain some antigen-binding function of the intact antigen-binding domain. Examples of antigen-binding fragments of an antibody include (1) a Fab fragment, a monovalent fragment having the VL, VH, CL and CH1 domains; (2) a F(ab')₂ fragment, a bivalent fragment having two Fab fragments linked by a disulfide bridge at the hinge region; (3) a Fd fragment having the two VH and CH1 domains; (4) a Fv fragment having the VL and VH domains of a single arm of an antibody, (5) a dAb fragment (Ward et al., (1989) Nature 341 :544-546), which has a VH domain; (6) an isolated complementarity determining region (CDR), and (7) a single chain Fv (scFv). Although the two domains of the Fv fragment, VL and VH are coded for by separate genes, they can be joined, using recombinant methods, by a synthetic linker that enables them to be made as a single protein chain in which the VL and VH regions pair to form monovalent molecules (known as single chain Fv (scFv); see e.g., Huston et al. (1988) Proc. Natl. Acad. Sci USA 85:5879-5883). These antibody fragments are obtained using conventional techniques known to those with skill in the art, and the fragments are evaluated for function in the same manner as are intact antibodies.

In the event that a (synthetic) linker is used, this linker is preferably of a length and sequence sufficient to ensure that each of the first and second domains can, independently from one another, retain their differential binding specificities. Most preferably and as documented in

the appended examples, the antibody construct of the invention is a “bispecific single chain antibody construct”, more preferably a bispecific single chain Fv (scFv). Bispecific single chain molecules are known in the art and are described in WO 99/54440, Mack, J. Immunol. (1997), 158, 3965-3970, Mack, PNAS, (1995), 92, 7021-7025, Kufer, Cancer Immunol. Immunother., (1997), 45, 193-197, Löffler, Blood, (2000), 95, 6, 2098-2103, Brühl, Immunol., (2001), 166, 2420-2426, Kipriyanov, J. Mol. Biol., (1999), 293, 41-56. One example of a CD33 targeting compound in connection with the present invention, which is a bispecific single chain molecule is AMG330, which has also been used in the appended examples.

The said variable domains comprised in the herein described antibody constructs may be connected by additional linker sequences. The term “peptide linker” defines in accordance with the present invention an amino acid sequence by which the amino acid sequences of the first domain and the second domain of the antibody construct of the invention are linked with each other. An essential technical feature of such peptide linker is that said peptide linker does not comprise any polymerization activity. Preferred amino acid residues for a peptide linker include Gly, Ser and Thr are characterized by a length between 5 and 25 amino acid residues. Among the suitable peptide linkers are those described in U.S. Patents 4,751,180 and 4,935,233 or WO 88/09344. A preferred embodiment of a peptide linker is characterized by the amino acid sequence Gly-Gly-Gly-Gly-Ser, i.e. Gly₄Ser, or polymers thereof, i.e. (Gly₄Ser)_x, where x is an integer 1 or greater. The characteristics of said peptide linker, which comprise the absence of the promotion of secondary structures are known in the art and described e.g. in Dall’Acqua et al. (Biochem. (1998) 37, 9266-9273), Cheadle et al. (Mol Immunol (1992) 29, 21-30) and Raag and Whitlow (FASEB (1995) 9(1), 73-80). Peptide linkers which also do not promote any secondary structures are preferred. The linkage of said domains to each other can be provided by, e.g. genetic engineering, as described in the examples. Methods for preparing fused and operatively linked bispecific single chain constructs and expressing them in mammalian cells or bacteria are well-known in the art (e.g. WO 99/54440 or Sambrook et al., Molecular Cloning: A Laboratory Manual, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, New York, 2001).

For peptide linkers, which connect the at least two binding domains in the antibody construct of the invention peptide linkers are preferred which comprise only a few number of amino acid residues, e.g. 12 amino acid residues or less. Thus, peptide linker of 12, 11, 10, 9, 8, 7, 6 or 5 amino acid residues are preferred. An envisaged peptide linker with less than 5 amino acids comprises 4, 3, 2 or one amino acid(s) wherein Gly-rich linkers are preferred. A particularly preferred “single” amino acid in context of said “peptide linker” is Gly. Accordingly, said peptide linker may consist of the single amino acid Gly.

The term "monoclonal antibody" as used herein refers to an antibody obtained from a population of substantially homogeneous antibodies, i.e., the individual antibodies comprising the population are identical except for possible naturally occurring mutations and/or post- translation modifications (e.g., isomerizations, amidations) that may be present in minor amounts. Monoclonal antibodies are highly specific, being directed against a single antigenic site. Furthermore, in contrast to conventional (polyclonal) antibody preparations which typically include different antibodies directed against different determinants (epitopes), each monoclonal antibody is directed against a single determinant on the antigen. In addition to their specificity, the monoclonal antibodies are advantageous in that they are synthesized by the hybridoma culture, uncontaminated by other immunoglobulins. The modifier "monoclonal" indicates the character of the antibody as being obtained from a substantially homogeneous population of antibodies, and is not to be construed as requiring production of the antibody by any particular method. For example, the monoclonal antibodies to be used in accordance with the present invention may be made by the hybridoma method first described by Kohler *et al.*, Nature, 256: 495 (1975), or may be made by recombinant DNA methods (see, e.g., U. S. Patent No. 4,816,567). The "monoclonal antibodies" may also be isolated from phage antibody libraries using the techniques described in Clackson *et al.*, Nature, 352: 624-628 (1991) and Marks *et al.*, J. Mol. Biol., 222: 581-597 (1991), for example.

The term "human antibody" includes antibodies having variable and constant regions corresponding substantially to human germline immunoglobulin sequences known in the art, including, for example, those described by Kabat *et al.* (See Kabat *et al.* (1991) loc. cit.). The human antibodies of the invention may include amino acid residues not encoded by human germline immunoglobulin sequences (e.g., mutations introduced by random or site-specific mutagenesis in vitro or by somatic mutation in vivo), for example in the CDRs, and in particular, CDR3. The human antibody can have at least one, two, three, four, five, or more positions replaced with an amino acid residue that is not encoded by the human germline immunoglobulin sequence. It is emphasized that the definition of human antibodies as used herein also contemplates fully human antibodies, which include only non-artificially and/or genetically altered human sequences of antibodies as those can be derived by using technologies using systems such as the Xenomice.

Examples of "antibody variants" include humanized variants of non- human antibodies, "affinity matured" antibodies (see, e.g. Hawkins *et al.* J. Mol. Biol. 254, 889-896 (1992) and Lowman *et al.*, Biochemistry 30, 10832- 10837 (1991)) and antibody mutants with altered effector function (s) (see, e.g., US Patent 5, 648, 260, Kontermann and Dübel (2010), loc. cit. and Little(2009), loc. cit.).

As used herein, "in vitro generated antibody" refers to an antibody where all or part of the variable region (e.g., at least one CDR) is generated in a non-immune cell selection (e.g., an in vitro phage display, protein chip or any other method in which candidate sequences can be tested for their ability to bind to an antigen). This term thus preferably excludes sequences generated by genomic rearrangement in an immune cell.

The pairing of a VH and VL together forms a single antigen-binding site. The CH domain most proximal to VH is designated as CH1. Each L chain is linked to an H chain by one covalent disulfide bond, while the two H chains are linked to each other by one or more disulfide bonds depending on the H chain isotype. The VH and VL domains consist of four regions of relatively conserved sequences called framework regions (FR1, FR2, FR3, and FR4), which form a scaffold for three regions of hypervariable sequences (complementarity determining regions, CDRs). The CDRs contain most of the residues responsible for specific interactions of the antibody with the antigen. CDRs are referred to as CDR 1, CDR2, and CDR3. Accordingly, CDR constituents on the heavy chain are referred to as H1, H2, and H3, while CDR constituents on the light chain are referred to as L1, L2, and L3.

The term "variable" refers to the portions of the immunoglobulin domains that exhibit variability in their sequence and that are involved in determining the specificity and binding affinity of a particular antibody (i.e., the "variable domain(s)"). Variability is not evenly distributed throughout the variable domains of antibodies; it is concentrated in sub-domains of each of the heavy and light chain variable regions. These sub-domains are called "hypervariable" regions or "complementarity determining regions" (CDRs). The more conserved (i.e., non-hypervariable) portions of the variable domains are called the "framework" regions (FRM). The variable domains of naturally occurring heavy and light chains each comprise four FRM regions, largely adopting a β -sheet configuration, connected by three hypervariable regions, which form loops connecting, and in some cases forming part of, the β -sheet structure. The hypervariable regions in each chain are held together in close proximity by the FRM and, with the hypervariable regions from the other chain, contribute to the formation of the antigen-binding site (see Kabat *et al.*, loc. cit.). The constant domains are not directly involved in antigen binding, but exhibit various effector functions, such as, for example, antibody-dependent, cell-mediated cytotoxicity and complement activation.

The terms "CDR", and its plural "CDRs", refer to a complementarity determining region (CDR) of which three make up the binding character of a light chain variable region (CDRL1, CDRL2 and CDRL3) and three make up the binding character of a heavy chain variable region (CDRH1, CDRH2 and CDRH3). CDRs contribute to the functional activity of an antibody molecule and are separated by amino acid sequences that comprise scaffolding or

framework regions. The exact definitional CDR boundaries and lengths are subject to different classification and numbering systems. CDRs may therefore be referred to by Kabat, Chothia, contact or any other boundary definitions, including the numbering system described herein. Despite differing boundaries, each of these systems has some degree of overlap in what constitutes the so called "hypervariable regions" within the variable sequences. CDR definitions according to these systems may therefore differ in length and boundary areas with respect to the adjacent framework region. See for example Kabat, Chothia, and/or MacCallum (Kabat *et al.*, loc. cit.; Chothia *et al.*, J. Mol. Biol, 1987, 196: 901; and MacCallum *et al.*, J. Mol. Biol, 1996, 262: 732). However, the numbering in accordance with the so-called Kabat system is preferred. The CDR3 of the light chain and, particularly, CDR3 of the heavy chain may constitute the most important determinants in antigen binding within the light and heavy chain variable regions. In some antibody constructs, the heavy chain CDR3 appears to constitute the major area of contact between the antigen and the antibody. In vitro selection schemes in which CDR3 alone is varied can be used to vary the binding properties of an antibody or determine which residues contribute to the binding of an antigen.

"Consisting essentially of" means that the amino acid sequence can vary by about 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, or 15% relative to the recited SEQ ID NO: sequence and still retain biological activity, as described herein.

In some embodiments, the binding molecules described herein are isolated proteins or substantially pure proteins. An "isolated" protein is unaccompanied by at least some of the material with which it is normally associated in its natural state, for example constituting at least about 5%, or at least about 50% by weight of the total protein in a given sample. It is understood that the isolated protein may constitute from 5 to 99.9% by weight of the total protein content depending on the circumstances. For example, the protein may be made at a significantly higher concentration through the use of an inducible promoter or high expression promoter, such that the protein is made at increased concentration levels. The definition includes the production of an antigen binding protein in a wide variety of organisms and/or host cells that are known in the art.

For amino acid sequences, sequence identity and/or similarity is determined by using standard techniques known in the art, including, but not limited to, the local sequence identity algorithm of Smith and Waterman, 1981, *Adv. Appl. Math.* 2:482, the sequence identity alignment algorithm of Needleman and Wunsch, 1970, *J. Mol. Biol.* 48:443, the search for similarity method of Pearson and Lipman, 1988, *Proc. Nat. Acad. Sci. U.S.A.* 85:2444, computerized implementations of these algorithms (GAP, BESTFIT, FASTA, and TFASTA in the Wisconsin Genetics Software Package, Genetics Computer Group, 575

Science Drive, Madison, Wis.), the Best Fit sequence program described by Devereux *et al.*, 1984, *Nucl. Acid Res.* 12:387-395, preferably using the default settings, or by inspection. Preferably, percent identity is calculated by FastDB based upon the following parameters: mismatch penalty of 1; gap penalty of 1; gap size penalty of 0.33; and joining penalty of 30,
5 "Current Methods in Sequence Comparison and Analysis," Macromolecule Sequencing and Synthesis, Selected Methods and Applications, pp 127-149 (1988), Alan R. Liss, Inc.

An example of a useful algorithm is PILEUP. PILEUP creates a multiple sequence alignment from a group of related sequences using progressive, pairwise alignments. It can also plot a tree showing the clustering relationships used to create the alignment. PILEUP uses a
10 simplification of the progressive alignment method of Feng & Doolittle, 1987, *J. Mol. Evol.* 35:351-360; the method is similar to that described by Higgins and Sharp, 1989, *CABIOS* 5:151-153. Useful PILEUP parameters including a default gap weight of 3.00, a default gap length weight of 0.10, and weighted end gaps.

Another example of a useful algorithm is the BLAST algorithm, described in: Altschul *et al.*,
15 1990, *J. Mol. Biol.* 215:403-410; Altschul *et al.*, 1997, *Nucleic Acids Res.* 25:3389-3402; and Karin *et al.*, 1993, *Proc. Natl. Acad. Sci. U.S.A.* 90:5873-5787. A particularly useful BLAST program is the WU-BLAST-2 program which was obtained from Altschul *et al.*, 1996, *Methods in Enzymology* 266:460-480. WU-BLAST-2 uses several search parameters, most of which are set to the default values. The adjustable parameters are set with the following
20 values: overlap span=1, overlap fraction=0.125, word threshold (T)=11. The HSP S and HSP S2 parameters are dynamic values and are established by the program itself depending upon the composition of the particular sequence and composition of the particular database against which the sequence of interest is being searched; however, the values may be adjusted to increase sensitivity.

An additional useful algorithm is gapped BLAST as reported by Altschul *et al.*, 1993, *Nucl. Acids Res.* 25:3389-3402. Gapped BLAST uses BLOSUM-62 substitution scores; threshold
25 T parameter set to 9; the two-hit method to trigger ungapped extensions, charges gap lengths of k a cost of 10+k; Xu set to 16, and Xg set to 40 for database search stage and to 67 for the output stage of the algorithms. Gapped alignments are triggered by a score
30 corresponding to about 22 bits.

Generally, the amino acid homology, similarity, or identity between individual variant CDRs are at least 80% to the sequences depicted herein, and more typically with preferably increasing homologies or identities of at least 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, and almost 100%. In a similar manner, "percent (%) nucleic acid sequence
35 identity" with respect to the nucleic acid sequence of the binding proteins identified herein is

defined as the percentage of nucleotide residues in a candidate sequence that are identical with the nucleotide residues in the coding sequence of the antigen binding protein. A specific method utilizes the BLASTN module of WU-BLAST-2 set to the default parameters, with overlap span and overlap fraction set to 1 and 0.125, respectively.

- 5 Generally, the nucleic acid sequence homology, similarity, or identity between the nucleotide sequences encoding individual variant CDRs and the nucleotide sequences depicted herein are at least 80%, and more typically with preferably increasing homologies or identities of at least 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, or 99%, and almost 100%.
- 10 Thus, a "variant CDR" is one with the specified homology, similarity, or identity to the parent CDR of the invention, and shares biological function, including, but not limited to, at least 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, or 99% of the specificity and/or activity of the parent CDR.

- While the site or region for introducing an amino acid sequence variation is predetermined,
- 15 the mutation *per se* need not be predetermined. For example, in order to optimize the performance of a mutation at a given site, random mutagenesis may be conducted at the target codon or region and the expressed antigen binding protein CDR variants screened for the optimal combination of desired activity. Techniques for making substitution mutations at predetermined sites in DNA having a known sequence are well known, for example, M13
- 20 primer mutagenesis and PCR mutagenesis. Screening of the mutants is done using assays of antigen binding protein activities, such as binding to an elected a cell surface molecule on a target cell.

- The term "amino acid" or "amino acid residue" typically refers to an amino acid having its art recognized definition such as an amino acid selected from the group consisting of: alanine
- 25 (Ala or A); arginine (Arg or R); asparagine (Asn or N); aspartic acid (Asp or D); cysteine (Cys or C); glutamine (Gln or Q); glutamic acid (Glu or E); glycine (Gly or G); histidine (His or H); isoleucine (He or I); leucine (Leu or L); lysine (Lys or K); methionine (Met or M); phenylalanine (Phe or F); pro line (Pro or P); serine (Ser or S); threonine (Thr or T); tryptophan (Trp or W); tyrosine (Tyr or Y); and valine (Val or V), although modified, synthetic,
- 30 or rare amino acids may be used as desired. Generally, amino acids can be grouped as having a nonpolar side chain (e.g., Ala, Cys, He, Leu, Met, Phe, Pro, Val); a negatively charged side chain (e.g., Asp, Glu); a positively charged sidechain (e.g., Arg, His, Lys); or an uncharged polar side chain (e.g., Asn, Cys, Gln, Gly, His, Met, Phe, Ser, Thr, Trp, and Tyr).
- The term "hypervariable region" (also known as "complementarity determining regions" or
- 35 CDRs) when used herein refers to the amino acid residues of an antibody which are (usually

three or four short regions of extreme sequence variability) within the V-region domain of an immunoglobulin which form the antigen-binding site and are the main determinants of antigen specificity. There are at least two methods for identifying the CDR residues: (1) An approach based on cross-species sequence variability (i. e., Kabat *et al.*, loc. cit.); and (2) An approach based on crystallographic studies of antigen-antibody complexes (Chothia, C. *et al.*, J. Mol. Biol. 196: 901-917 (1987)). However, to the extent that two residue identification techniques define regions of overlapping, but not identical regions, they can be combined to define a hybrid CDR. However, in general, the CDR residues are preferably identified in accordance with the so-called Kabat (numbering) system.

The term "framework region" refers to the art-recognized portions of an antibody variable region that exist between the more divergent (i.e., hypervariable) CDRs. Such framework regions are typically referred to as frameworks 1 through 4 (FR1, FR2, FR3, and FR4) and provide a scaffold for the presentation of the six CDRs (three from the heavy chain and three from the light chain) in three dimensional space, to form an antigen-binding surface.

Typically, CDRs form a loop structure that can be classified as a canonical structure. The term "canonical structure" refers to the main chain conformation that is adopted by the antigen binding (CDR) loops. From comparative structural studies, it has been found that five of the six antigen binding loops have only a limited repertoire of available conformations. Each canonical structure can be characterized by the torsion angles of the polypeptide backbone. Correspondent loops between antibodies may, therefore, have very similar three dimensional structures, despite high amino acid sequence variability in most parts of the loops (Chothia and Lesk, J. Mol. Biol., 1987, 196: 901; Chothia *et al.*, Nature, 1989, 342: 877; Martin and Thornton, J. Mol. Biol, 1996, 263: 800, each of which is incorporated by reference in its entirety). Furthermore, there is a relationship between the adopted loop structure and the amino acid sequences surrounding it. The conformation of a particular canonical class is determined by the length of the loop and the amino acid residues residing at key positions within the loop, as well as within the conserved framework (i.e., outside of the loop). Assignment to a particular canonical class can therefore be made based on the presence of these key amino acid residues. The term "canonical structure" may also include considerations as to the linear sequence of the antibody, for example, as catalogued by Kabat (Kabat *et al.*, loc. cit.). The Kabat numbering scheme (system) is a widely adopted standard for numbering the amino acid residues of an antibody variable domain in a consistent manner and is the preferred scheme applied in the present invention as also mentioned elsewhere herein. Additional structural considerations can also be used to determine the canonical structure of an antibody. For example, those differences not fully reflected by Kabat numbering can be described by the numbering system of Chothia *et al* and/or revealed by other techniques, for example, crystallography and two or three-

dimensional computational modeling. Accordingly, a given antibody sequence may be placed into a canonical class which allows for, among other things, identifying appropriate chassis sequences (e.g., based on a desire to include a variety of canonical structures in a library). Kabat numbering of antibody amino acid sequences and structural considerations as described by Chothia *et al.*, loc. cit. and their implications for construing canonical aspects of antibody structure, are described in the literature.

CDR3 is typically the greatest source of molecular diversity within the antibody-binding site. H3, for example, can be as short as two amino acid residues or greater than 26 amino acids. The subunit structures and three-dimensional configurations of different classes of immunoglobulins are well known in the art. For a review of the antibody structure, see Antibodies: A Laboratory Manual, Cold Spring Harbor Laboratory, eds. Harlow *et al.*, 1988. One of skill in the art will recognize that each subunit structure, e.g., a CH, VH, CL, VL, CDR, FR structure, comprises active fragments, e.g., the portion of the VH, VL, or CDR subunit that binds to the antigen, i.e., the antigen-binding fragment, or, e.g., the portion of the CH subunit that binds to and/or activates, e.g., an Fc receptor and/or complement. The CDRs typically refer to the Kabat CDRs, as described in Sequences of Proteins of immunological Interest, US Department of Health and Human Services (1991), eds. Kabat *et al.* Another standard for characterizing the antigen binding site is to refer to the hypervariable loops as described by Chothia. See, e.g., Chothia, *et al.* (1987; J. Mol. Biol. 227:799-817); and Tomlinson *et al.* (1995) EMBO J. 14: 4628-4638. Still another standard is the AbM definition used by Oxford Molecular's AbM antibody modeling software. See, generally, e.g., Protein Sequence and Structure Analysis of Antibody Variable Domains. In: Antibody Engineering Lab Manual (Ed.: Duebel, S. and Kontermann, R., Springer-Verlag, Heidelberg). Embodiments described with respect to Kabat CDRs can alternatively be implemented using similar described relationships with respect to Chothia hypervariable loops or to the AbM-defined loops.

The sequence of antibody genes after assembly and somatic mutation is highly varied, and these varied genes are estimated to encode 10^{10} different antibody molecules (Immunoglobulin Genes, 2nd ed., eds. Jonio *et al.*, Academic Press, San Diego, CA, 1995). Accordingly, the immune system provides a repertoire of immunoglobulins. The term "repertoire" refers to at least one nucleotide sequence derived wholly or partially from at least one sequence encoding at least one immunoglobulin. The sequence(s) may be generated by rearrangement in vivo of the V, D, and J segments of heavy chains, and the V and J segments of light chains. Alternatively, the sequence(s) can be generated from a cell in response to which rearrangement occurs, e.g., in vitro stimulation. Alternatively, part or all of the sequence(s) may be obtained by DNA splicing, nucleotide synthesis, mutagenesis, and other methods, see, e.g., U.S. Patent 5,565,332. A repertoire may include only one

sequence or may include a plurality of sequences, including ones in a genetically diverse collection.

It is also envisaged that the CD33 targeting compound described herein has, apart from its function to bind to the cell surface molecule CD33 on a target cell and CD3 on the cell surface of a T cell, may have an additional function. In this format, the compound is a multifunctional compound by targeting cells through binding to CD33 on the cell surface of a target cell, mediating cytotoxic T cell activity through CD3 binding and providing a further function such as a fully functional Fc constant domain mediating antibody-dependent cellular cytotoxicity through recruitment of effector cells like NK cells, a half life extending domain such as an albumin binding domain or a modified Fc constant domain lacking antibody-dependent cellular cytotoxicity but extending the molecular weight of the compound, mediation of a label (fluorescent etc.), a therapeutic agent such as, e.g. a toxin or radionuclide, and/or means to enhance serum half-life, etc.

The term "epitope" refers to a site on an antigen to which a binding domain, such as an antibody or immunoglobulin or derivative or fragment of an antibody or of an immunoglobulin, specifically binds. An "epitope" is antigenic and thus the term epitope is sometimes also referred to herein as "antigenic structure" or "antigenic determinant". Thus, the binding domain is an "antigen-interaction-site". Said binding/interaction is also understood to define a "specific recognition". In one example, said binding domain which (specifically) binds to / interacts with a given target epitope of CD33 on a cell surface molecule on a target cell or CD3 is an antibody or immunoglobulin, and said binding domain is a VH and/or VL region of an antibody or of an immunoglobulin.

"Epitopes" can be formed both by contiguous amino acids or non-contiguous amino acids juxtaposed by tertiary folding of a protein. A "linear epitope" is an epitope where an amino acid primary sequence comprises the recognized epitope. A linear epitope typically includes at least 3 or at least 4, and more usually, at least 5 or at least 6 or at least 7, for example, about 8 to about 10 amino acids in a unique sequence.

A "conformational epitope", in contrast to a linear epitope, is an epitope wherein the primary sequence of the amino acids comprising the epitope is not the sole defining component of the epitope recognized (e.g., an epitope wherein the primary sequence of amino acids is not necessarily recognized by the binding domain). Typically a conformational epitope comprises an increased number of amino acids relative to a linear epitope. With regard to recognition of conformational epitopes, the binding domain recognizes a three-dimensional structure of the antigen, preferably a peptide or protein or fragment thereof (in the context of the present invention, the antigen for one of the binding domains is comprised within a cell surface molecule on a target cell). For example, when a protein molecule folds to form a

three-dimensional structure, certain amino acids and/or the polypeptide backbone forming the conformational epitope become juxtaposed enabling the antibody to recognize the epitope. Methods of determining the conformation of epitopes include, but are not limited to, x-ray crystallography, two-dimensional nuclear magnetic resonance (2D-NMR) spectroscopy and site-directed spin labelling and electron paramagnetic resonance (EPR) spectroscopy. Moreover, the provided examples describe a further method to characterize a given binding domain by way of binning, which includes a test whether the given binding domain binds to one or more epitope cluster(s) of a given protein, in particular a cell surface molecule on a target cell.

As used herein, the term "epitope cluster" denotes the entirety of epitopes lying in a defined contiguous stretch of an antigen. An epitope cluster can comprise one, two or more epitopes. The concept of epitope cluster is also used in the characterization of the features of the binding molecules of the invention.

The terms "(capable of) binding to", "specifically recognizing", "directed to" and "reacting with" mean in accordance with this invention that a binding domain is capable of specifically interacting with one or more, preferably at least two, more preferably at least three and most preferably at least four amino acids of an epitope.

As used herein, the terms "specifically interacting", "specifically binding" or "specifically bind(s)" mean that a binding domain exhibits appreciable affinity for a particular protein or antigen and, generally, does not exhibit significant reactivity with proteins or antigens other than CD33 or CD3. "Appreciable affinity" includes binding with an affinity of about 10^{-6} M (KD) or stronger. Preferably, binding is considered specific when binding affinity is about 10^{-12} to 10^{-8} M, 10^{-12} to 10^{-9} M, 10^{-12} to 10^{-10} M, 10^{-11} to 10^{-8} M, preferably of about 10^{-11} to 10^{-9} M. Whether a binding domain specifically reacts with or binds to a target can be tested readily by, *inter alia*, comparing the reaction of said binding domain with a target protein or antigen with the reaction of said binding domain with proteins or antigens other than CD33 or CD3. Preferably, a binding domain of the invention does not essentially bind or is not capable of binding to proteins or antigens other than CD33 or CD3 (i.e. the first binding domain is not capable of binding to proteins other than CD33 and the second binding domain is not capable of binding to proteins other than CD3).

The term "does not essentially bind", or "is not capable of binding" means that a binding domain of the present invention does not bind another protein or antigen other than CD33 or CD3, i.e., does not show reactivity of more than 30%, preferably not more than 20%, more preferably not more than 10%, particularly preferably not more than 9%, 8%, 7%, 6% or 5%

with proteins or antigens other than CD33 or CD3, whereby binding to CD33 or CD3, respectively, is set to be 100%.

Specific binding is believed to be effected by specific motifs in the amino acid sequence of the binding domain and the antigen. Thus, binding is achieved as a result of their primary, secondary and/or tertiary structure as well as the result of secondary modifications of said structures. The specific interaction of the antigen-interaction-site with its specific antigen may result in a simple binding of said site to the antigen. Moreover, the specific interaction of the antigen-interaction-site with its specific antigen may alternatively or additionally result in the initiation of a signal, e.g. due to the induction of a change of the conformation of the antigen, an oligomerization of the antigen, etc.

Proteins (including fragments thereof, preferably biologically active fragments, and peptides, usually having less than 30 amino acids) comprise one or more amino acids coupled to each other via a covalent peptide bond (resulting in a chain of amino acids). The term "polypeptide" as used herein describes a group of molecules, which consist of more than 30 amino acids. Polypeptides may further form multimers such as dimers, trimers and higher oligomers, i.e. consisting of more than one polypeptide molecule. Polypeptide molecules forming such dimers, trimers etc. may be identical or non-identical. The corresponding higher order structures of such multimers are, consequently, termed homo- or heterodimers, homo- or heterotrimers etc. An example for a heteromultimer is an antibody molecule, which, in its naturally occurring form, consists of two identical light polypeptide chains and two identical heavy polypeptide chains. The terms "polypeptide" and "protein" also refer to naturally modified polypeptides/proteins wherein the modification is effected e.g. by post-translational modifications like glycosylation, acetylation, phosphorylation and the like. A "polypeptide" when referred to herein may also be chemically modified such as pegylated. Such modifications are well known in the art.

"Isolated" when used to describe the CD33 targeting compound disclosed herein, means a compound that has been identified, separated and/or recovered from a component of its production environment. Preferably, the isolated compound is free of association with all other components from its production environment. Contaminant components of its production environment, such as that resulting from recombinant transfected cells, are materials that would typically interfere with diagnostic or therapeutic uses for the polypeptide, and may include enzymes, hormones, and other proteinaceous or non-proteinaceous solutes. In preferred embodiments, the compound will be purified (1) to a degree sufficient to obtain at least 15 residues of N-terminal or internal amino acid sequence by use of a spinning cup sequenator, or (2) to homogeneity by SDS-PAGE under non-

reducing or reducing conditions using Coomassie blue or, preferably, silver stain. Ordinarily, however, an isolated compound will be prepared by at least one purification step.

Amino acid sequence modifications of the CD33 targeting compound described herein are contemplated. For example, it may be desirable to improve the binding affinity and/or other biological properties of the compound. Amino acid sequence variants of the CD33 targeting compounds are prepared by introducing appropriate nucleotide changes into the compounds nucleic acid, or by peptide synthesis.

Such modifications include, for example, deletions from, and/or insertions into, and/or substitutions of, residues within the amino acid sequences of the compound. Any combination of deletion, insertion, and substitution is made to arrive at the final construct, provided that the final construct possesses the desired characteristics. The amino acid changes also may alter post-translational processes of the compound, such as changing the number or position of glycosylation sites. Preferably, 1, 2, 3, 4, 5, 6, 7, 8, 9, or 10 amino acids may be substituted in a CDR, while 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, or 25 amino acids may be substituted in the framework regions (FRs). The substitutions are preferably conservative substitutions as described herein. Additionally or alternatively, 1, 2, 3, 4, 5, or 6 amino acids may be inserted or deleted in each of the CDRs (of course, dependent on their length), while 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, or 25 amino acids may be inserted or deleted in each of the FRs.

A useful method for identification of certain residues or regions of the CD33 targeting compounds that are preferred locations for mutagenesis is called "alanine scanning mutagenesis" as described by Cunningham and Wells in *Science*, 244: 1081-1085 (1989). Here, a residue or group of target residues within the compound is/are identified (e.g. charged residues such as arg, asp, his, lys, and glu) and replaced by a neutral or negatively charged amino acid (most preferably alanine or polyalanine) to affect the interaction of the amino acids with the epitope.

Those amino acid locations demonstrating functional sensitivity to the substitutions then are refined by introducing further or other variants at, or for, the sites of substitution. Thus, while the site for introducing an amino acid sequence variation is predetermined, the nature of the mutation *per se* needs not to be predetermined. For example, to analyze the performance of a mutation at a given site, ala scanning or random mutagenesis is conducted at a target codon or region and the expressed compound variants are screened for the desired activity.

Preferably, amino acid sequence insertions include amino- and/or carboxyl-terminal fusions ranging in length from 1, 2, 3, 4, 5, 6, 7, 8, 9 or 10 residues to polypeptides containing a hundred or more residues, as well as intrasequence insertions of single or multiple amino

acid residues. An insertional variant of the CD33 targeting compound includes the fusion to the N-or C-terminus of the antibody to an enzyme or a fusion to a polypeptide which increases the serum half-life of the antibody.

Another type of variant is an amino acid substitution variant. These variants have preferably at least 1, 2, 3, 4, 5, 6, 7, 8, 9 or 10 amino acid residues in the compound replaced by a different residue. The sites of greatest interest for substitutional mutagenesis include the CDRs of the heavy and/or light chain, in particular the hypervariable regions, but FR alterations in the heavy and/or light chain are also contemplated.

For example, if a CDR sequence encompasses 6 amino acids, it is envisaged that one, two or three of these amino acids are substituted. Similarly, if a CDR sequence encompasses 15 amino acids it is envisaged that one, two, three, four, five or six of these amino acids are substituted.

Generally, if amino acids are substituted in one or more or all of the CDRs of the heavy and/or light chain, it is preferred that the then-obtained "substituted" sequence is at least 60%, more preferably 65%, even more preferably 70%, particularly preferably 75%, more particularly preferably 80% identical to the "original" CDR sequence. This means that it is dependent of the length of the CDR to which degree it is identical to the "substituted" sequence. For example, a CDR having 5 amino acids is preferably 80% identical to its substituted sequence in order to have at least one amino acid substituted. Accordingly, the CDRs of the CD33 targeting compound may have different degrees of identity to their substituted sequences, e.g., CDRL1 may have 80%, while CDRL3 may have 90%.

Preferred substitutions (or replacements) are conservative substitutions. However, any substitution (including non-conservative substitution or one or more from the "exemplary substitutions" listed in Table 1, below) is envisaged as long as the CD33 targeting compound retains its capability to bind to CD33 via the first binding domain and to CD3 epsilon via the second binding domain and/or its CDRs have an identity to the then substituted sequence (at least 60%, more preferably 65%, even more preferably 70%, particularly preferably 75%, more particularly preferably 80% identical to the "original" CDR sequence).

Conservative substitutions are shown in Table 1 under the heading of "preferred substitutions". If such substitutions result in a change in biological activity, then more substantial changes, denominated "exemplary substitutions" in Table 1, or as further described below in reference to amino acid classes, may be introduced and the products screened for a desired characteristic.

Table 1: Amino Acid Substitutions

Original	Exemplary Substitutions	Preferred Substitutions
Ala (A)	val, leu, ile	val
Arg (R)	lys, gln, asn	lys
Asn (N)	gln, his, asp, lys, arg	gln
Asp (D)	glu, asn	glu
Cys (C)	ser, ala	ser
Gln (Q)	asn, glu	asn
Glu (E)	asp, gln	asp
Gly (G)	ala	ala
His (H)	asn, gln, lys, arg	arg
Ile (I)	leu, val, met, ala, phe	leu
Leu (L)	norleucine, ile, val, met, ala	ile
Lys (K)	arg, gln, asn	arg
Met (M)	leu, phe, ile	leu
Phe (F)	leu, val, ile, ala, tyr	tyr
Pro (P)	ala	ala
Ser (S)	thr	thr
Thr (T)	ser	ser
Trp (W)	tyr, phe	tyr
Tyr (Y)	trp, phe, thr, ser	phe
Val (V)	ile, leu, met, phe, ala	leu

Substantial modifications in the biological properties of the CD33 targeting compound described herein are accomplished by selecting substitutions that differ significantly in their effect on maintaining (a) the structure of the polypeptide backbone in the area of the substitution, for example, as a sheet or helical conformation, (b) the charge or hydrophobicity of the molecule at the target site, or (c) the bulk of the side chain. Naturally occurring residues are divided into groups based on common side-chain properties: (1) hydrophobic: norleucine, met, ala, val, leu, ile; (2) neutral hydrophilic: cys, ser, thr; (3) acidic: asp, glu; (4) basic: asn, gin, his, lys, arg; (5) residues that influence chain orientation: gly, pro; and (6) aromatic : trp, tyr, phe.

Non-conservative substitutions will entail exchanging a member of one of these classes for another class. Any cysteine residue not involved in maintaining the proper conformation of the CD33 targeting compound may be substituted, generally with serine, to improve the oxidative stability of the molecule and prevent aberrant crosslinking. Conversely, cysteine

bond(s) may be added to the compound in case the compound is an antibody to improve its stability (particularly where the antibody is an antibody fragment such as an Fv fragment).

A particularly preferred type of substitutional variant involves substituting one or more hypervariable region residues of a parent antibody (e. g. a humanized or human antibody).

5 Generally, the resulting variant(s) selected for further development will have improved biological properties relative to the parent antibody from which they are generated. A convenient way for generating such substitutional variants involves affinity maturation using phage display. Briefly, several hypervariable region sites (e. g. 6-7 sites) are mutated to generate all possible amino acid substitutions at each site. The antibody variants thus
10 generated are displayed in a monovalent fashion from filamentous phage particles as fusions to the gene III product of M13 packaged within each particle. The phage-displayed variants are then screened for their biological activity (e. g. binding affinity) as herein disclosed. In order to identify candidate hypervariable region sites for modification, alanine scanning mutagenesis can be performed to identify hypervariable region residues
15 contributing significantly to antigen binding. Alternatively, or additionally, it may be beneficial to analyze a crystal structure of the antigen-antibody complex to identify contact points between the binding domain and, e.g., human CD33 or CD3. Such contact residues and neighbouring residues are candidates for substitution according to the techniques elaborated herein. Once such variants are generated, the panel of variants is subjected to screening as
20 described herein and antibodies with superior properties in one or more relevant assays may be selected for further development.

Other modifications of the CD33 targeting compound are contemplated herein. For example, the CD33 targeting compound may be linked to one of a variety of non-proteinaceous polymers, e.g., polyethylene glycol, polypropylene glycol, polyoxyalkylenes, or copolymers of
25 polyethylene glycol and polypropylene glycol. The CD33 targeting compound may also be entrapped in microcapsules prepared, for example, by coacervation techniques or by interfacial polymerization (for example, hydroxymethylcellulose or gelatine-microcapsules and poly (methylmethacrylate) microcapsules, respectively), in colloidal drug delivery systems (for example, liposomes, albumin microspheres, microemulsions, nanoparticles and
30 nanocapsules), or in macroemulsions. Such techniques are disclosed in Remington's Pharmaceutical Sciences, 16th edition, Oslo, A., Ed., (1980).

The CD33 targeting compound disclosed herein may also be formulated as immuno-liposomes. A "liposome" is a small vesicle composed of various types of lipids, phospholipids and/or surfactant which is useful for delivery of a drug to a mammal. The components of the
35 liposome are commonly arranged in a bilayer formation, similar to the lipid arrangement of biological membranes. Liposomes containing the compound are prepared by methods

known in the art, such as described in Epstein *et al.*, Proc. Natl. Acad. Sci. USA, 82: 3688 (1985); Hwang *et al.*, Proc. Natl. Acad. Sci. USA, 77: 4030 (1980); US Pat. Nos. 4,485,045 and 4,544,545; and W0 97/38731 published October 23, 1997. Liposomes with enhanced circulation time are disclosed in US Patent No. 5,013, 556. Particularly useful liposomes can be generated by the reverse phase evaporation method with a lipid composition comprising phosphatidylcholine, cholesterol and PEG-derivatized phosphatidylethanolamine (PEG-PE). Liposomes are extruded through filters of defined pore size to yield liposomes with the desired diameter. Fab' fragments of the antibody of the present invention can be conjugated to the liposomes as described in Martin *et al.* J. Biol. Chem. 257: 286-288 (1982) via a disulfide interchange reaction. A chemotherapeutic agent is optionally contained within the liposome. See Gabizon *et al.* J. National Cancer Inst. 81 (19) 1484 (1989).

When using recombinant techniques, the CD33 targeting compound can be produced intracellularly, in the periplasmic space, or directly secreted into the medium. If the antibody construct is produced intracellularly, as a first step, the particulate debris, either host cells or lysed fragments, are removed, for example, by centrifugation or ultrafiltration. Carter *et al.*, Bio/Technology 10: 163-167 (1992) describe a procedure for isolating antibodies which are secreted to the periplasmic space of *E. coli*.

The CD33 targeting compounds prepared from the cells can be purified using, for example, hydroxylapatite chromatography, gel electrophoresis, dialysis, and affinity chromatography, with affinity chromatography being the preferred purification technique.

The CD33 targeting compounds described herein may be provided in form of a fusion protein comprising at least two binding domains, with or without peptide linkers (spacer peptides). Among the suitable peptide linkers are those described in U.S. Patents 4,751,180 and 4,935,233 or WO 88/09344.

Another method for preparing CD33 targeting compounds described herein in form of oligomeric antibody construct derivatives involves use of a leucine zipper. Leucine zipper domains are peptides that promote oligomerization of the proteins in which they are found. Leucine zippers were originally identified in several DNA-binding proteins (Landschulz *et al.*, 1988, *Science* 240:1759), and have since been found in a variety of different proteins. Among the known leucine zippers are naturally occurring peptides and derivatives thereof that dimerize or trimerize. Examples of leucine zipper domains suitable for producing soluble oligomeric proteins are described in PCT application WO 94/10308, and the leucine zipper derived from lung surfactant protein D (SPD) described in Hoppe *et al.*, 1994, *FEBS Letters* 344:191, hereby incorporated by reference. The use of a modified leucine zipper that allows for stable trimerization of a heterologous protein fused thereto is described in Fanslow *et al.*,

1994, *Semin. Immunol.* 6:267-78. In one approach, recombinant fusion proteins comprising a CD33 and a CD3 antibody fragment or derivative fused to a leucine zipper peptide are expressed in suitable host cells, and the soluble oligomeric CD33 and CD3 antibody fragments or derivatives that form are recovered from the culture supernatant.

- 5 Covalent modifications of antigen binding proteins are included within the scope of this invention, and are generally, but not always, done post-translationally. For example, several types of covalent modifications of the antigen binding protein are introduced into the molecule by reacting specific amino acid residues of the antigen binding protein with an organic derivatizing agent that is capable of reacting with selected side chains or the N- or
10 C-terminal residues.

Cysteiny l residues most commonly are reacted with α -haloacetates (and corresponding amines), such as chloroacetic acid or chloroacetamide, to give carboxymethyl or carboxyamidomethyl derivatives. Cysteiny l residues also are derivatized by reaction with
15 bromotrifluoroacetone, α -bromo- β -(5-imidazolyl)propionic acid, chloroacetyl phosphate, N-alkylmaleimides, 3-nitro-2-pyridyl disulfide, methyl 2-pyridyl disulfide, p-chloromercuribenzoate, 2-chloromercuri-4-nitrophenol, or chloro-7-nitrobenzo-2-oxa-1,3-diazole.

Histidyl residues are derivatized by reaction with diethylpyrocarbonate at pH 5.5-7.0 because this agent is relatively specific for the histidyl side chain. Para-bromophenacyl bromide also
20 is useful; the reaction is preferably performed in 0.1M sodium cacodylate at pH 6.0.

Lysiny l and amino terminal residues are reacted with succinic or other carboxylic acid anhydrides. Derivatization with these agents has the effect of reversing the charge of the lysiny l residues. Other suitable reagents for derivatizing alpha-amino-containing residues include imidoesters such as methyl picolinimidate; pyridoxal phosphate; pyridoxal;
25 chloroborohydride; trinitrobenzenesulfonic acid; O-methylisourea; 2,4-pentanedione; and transaminase-catalyzed reaction with glyoxylate.

Arginy l residues are modified by reaction with one or several conventional reagents, among them phenylglyoxal, 2,3-butanedione, 1,2-cyclohexanedione, and ninhydrin. Derivatization of arginine residues requires that the reaction be performed in alkaline conditions because of
30 the high pKa of the guanidine functional group. Furthermore, these reagents may react with the groups of lysine as well as the arginine epsilon-amino group.

The specific modification of tyrosyl residues may be made, with particular interest in introducing spectral labels into tyrosyl residues by reaction with aromatic diazonium compounds or tetranitromethane. Most commonly, N-acetylimidazole and tetranitromethane
35 are used to form O-acetyl tyrosyl species and 3-nitro derivatives, respectively. Tyrosyl

residues are iodinated using ^{125}I or ^{131}I to prepare labeled proteins for use in radioimmunoassay, the chloramine T method described above being suitable.

Carboxyl side groups (aspartyl or glutamyl) are selectively modified by reaction with carbodiimides ($\text{R}'\text{—N}=\text{C}=\text{N}\text{—R}'$), where R and R' are optionally different alkyl groups, such as 1-cyclohexyl-3-(2-morpholinyl-4-ethyl) carbodiimide or 1-ethyl-3-(4-azonia-4,4-dimethylpentyl) carbodiimide. Furthermore, aspartyl and glutamyl residues are converted to asparaginyl and glutaminyl residues by reaction with ammonium ions.

Derivatization with bifunctional agents is useful for crosslinking antigen binding proteins to a water-insoluble support matrix or surface for use in a variety of methods. Commonly used crosslinking agents include, e.g., 1,1-bis(diazoacetyl)-2-phenylethane, glutaraldehyde, N-hydroxysuccinimide esters, for example, esters with 4-azidosalicylic acid, homobifunctional imidoesters, including disuccinimidyl esters such as 3,3'-dithiobis(succinimidylpropionate), and bifunctional maleimides such as bis-N-maleimido-1,8-octane. Derivatizing agents such as methyl-3-[(p-azidophenyl)dithio]propioimide yield photoactivatable intermediates that are capable of forming crosslinks in the presence of light. Alternatively, reactive water-insoluble matrices such as cyanogen bromide-activated carbohydrates and the reactive substrates described in U.S. Pat. Nos. 3,969,287; 3,691,016; 4,195,128; 4,247,642; 4,229,537; and 4,330,440 are employed for protein immobilization.

Glutaminyl and asparaginyl residues are frequently deamidated to the corresponding glutamyl and aspartyl residues, respectively. Alternatively, these residues are deamidated under mildly acidic conditions. Either form of these residues falls within the scope of this invention.

Other modifications include hydroxylation of proline and lysine, phosphorylation of hydroxyl groups of seryl or threonyl residues, methylation of the α -amino groups of lysine, arginine, and histidine side chains (T. E. Creighton, *Proteins: Structure and Molecular Properties*, W. H. Freeman & Co., San Francisco, 1983, pp. 79-86), acetylation of the N-terminal amine, and amidation of any C-terminal carboxyl group.

Another type of covalent modification of the antigen binding protein included within the scope of this invention comprises altering the glycosylation pattern of the protein. As is known in the art, glycosylation patterns can depend on both the sequence of the protein (e.g., the presence or absence of particular glycosylation amino acid residues, discussed below), or the host cell or organism in which the protein is produced. Particular expression systems are discussed below.

Glycosylation of polypeptides is typically either N-linked or O-linked. N-linked refers to the attachment of the carbohydrate moiety to the side chain of an asparagine residue. The tri-

peptide sequences asparagine-X-serine and asparagine-X-threonine, where X is any amino acid except proline, are the recognition sequences for enzymatic attachment of the carbohydrate moiety to the asparagine side chain. Thus, the presence of either of these tri-peptide sequences in a polypeptide creates a potential glycosylation site. O-linked glycosylation refers to the attachment of one of the sugars N-acetylgalactosamine, galactose, or xylose, to a hydroxyamino acid, most commonly serine or threonine, although 5-hydroxyproline or 5-hydroxylysine may also be used.

Addition of glycosylation sites to the antigen binding protein is conveniently accomplished by altering the amino acid sequence such that it contains one or more of the above-described tri-peptide sequences (for N-linked glycosylation sites). The alteration may also be made by the addition of, or substitution by, one or more serine or threonine residues to the starting sequence (for O-linked glycosylation sites). For ease, the antigen binding protein amino acid sequence is preferably altered through changes at the DNA level, particularly by mutating the DNA encoding the target polypeptide at preselected bases such that codons are generated that will translate into the desired amino acids.

Another means of increasing the number of carbohydrate moieties on the antigen binding protein is by chemical or enzymatic coupling of glycosides to the protein. These procedures are advantageous in that they do not require production of the protein in a host cell that has glycosylation capabilities for N- and O-linked glycosylation. Depending on the coupling mode used, the sugar(s) may be attached to (a) arginine and histidine, (b) free carboxyl groups, (c) free sulfhydryl groups such as those of cysteine, (d) free hydroxyl groups such as those of serine, threonine, or hydroxyproline, (e) aromatic residues such as those of phenylalanine, tyrosine, or tryptophan, or (f) the amide group of glutamine. These methods are described in WO 87/05330 published Sep. 11, 1987, and in Aplin and Wriston, 1981, *CRC Crit. Rev. Biochem.*, pp. 259-306.

Removal of carbohydrate moieties present on the starting antigen binding protein may be accomplished chemically or enzymatically. Chemical deglycosylation requires exposure of the protein to the compound trifluoromethanesulfonic acid, or an equivalent compound. This treatment results in the cleavage of most or all sugars except the linking sugar (N-acetylglucosamine or N-acetylgalactosamine), while leaving the polypeptide intact. Chemical deglycosylation is described by Hakimuddin *et al.*, 1987, *Arch. Biochem. Biophys.* 259:52 and by Edge *et al.*, 1981, *Anal. Biochem.* 118:131. Enzymatic cleavage of carbohydrate moieties on polypeptides can be achieved by the use of a variety of endo- and exo-glycosidases as described by Thotakura *et al.*, 1987, *Meth. Enzymol.* 138:350. Glycosylation at potential glycosylation sites may be prevented by the use of the compound tunicamycin as

described by Duskin *et al.*, 1982, *J. Biol. Chem.* 257:3105. Tunicamycin blocks the formation of protein-N-glycoside linkages.

Another type of covalent modification of the antigen binding protein comprises linking the antigen binding protein to various non-proteinaceous polymers, including, but not limited to, various polyols such as polyethylene glycol, polypropylene glycol or polyoxyalkylenes, in the manner set forth in U.S. Pat. Nos. 4,640,835; 4,496,689; 4,301,144; 4,670,417; 4,791,192 or 4,179,337. In addition, as is known in the art, amino acid substitutions may be made in various positions within the antigen binding protein to facilitate the addition of polymers such as PEG.

- 10 In some embodiments, the covalent modification of the antigen binding proteins described herein comprises the addition of one or more labels.

The term "labelling group" means any detectable label. Examples of suitable labelling groups include, but are not limited to, the following: radioisotopes or radionuclides (*e.g.*, ^3H , ^{14}C , ^{15}N , ^{35}S , ^{90}Y , ^{99}Tc , ^{111}In , ^{125}I , ^{131}I), fluorescent groups (*e.g.*, FITC, rhodamine, lanthanide phosphors), enzymatic groups (*e.g.*, horseradish peroxidase, β -galactosidase, luciferase, alkaline phosphatase), chemiluminescent groups, biotinyl groups, or predetermined polypeptide epitopes recognized by a secondary reporter (*e.g.*, leucine zipper pair sequences, binding sites for secondary antibodies, metal binding domains, epitope tags). In some embodiments, the labelling group is coupled to the antigen binding protein *via* spacer arms of various lengths to reduce potential steric hindrance. Various methods for labelling proteins are known in the art and may be used in performing the present invention.

In general, labels fall into a variety of classes, depending on the assay in which they are to be detected: a) isotopic labels, which may be radioactive or heavy isotopes; b) magnetic labels (*e.g.*, magnetic particles); c) redox active moieties; d) optical dyes; enzymatic groups (*e.g.* horseradish peroxidase, β -galactosidase, luciferase, alkaline phosphatase); e) biotinylated groups; and f) predetermined polypeptide epitopes recognized by a secondary reporter (*e.g.*, leucine zipper pair sequences, binding sites for secondary antibodies, metal binding domains, epitope tags, etc.). In some embodiments, the labelling group is coupled to the antigen binding protein *via* spacer arms of various lengths to reduce potential steric hindrance. Various methods for labelling proteins are known in the art and may be used in performing the present invention.

Specific labels include optical dyes, including, but not limited to, chromophores, phosphors and fluorophores, with the latter being specific in many instances. Fluorophores can be either "small molecule" fluo- res, or proteinaceous fluo- res.

By "fluorescent label" is meant any molecule that may be detected via its inherent fluorescent properties. Suitable fluorescent labels include, but are not limited to, fluorescein, rhodamine, tetramethylrhodamine, eosin, erythrosin, coumarin, methyl-coumarins, pyrene, Malacite green, stilbene, Lucifer Yellow, Cascade BlueJ, Texas Red, IAEDANS, EDANS, BODIPY FL, LC Red 640, Cy 5, Cy 5.5, LC Red 705, Oregon green, the Alexa-Fluor dyes (Alexa Fluor 350, Alexa Fluor 430, Alexa Fluor 488, Alexa Fluor 546, Alexa Fluor 568, Alexa Fluor 594, Alexa Fluor 633, Alexa Fluor 660, Alexa Fluor 680), Cascade Blue, Cascade Yellow and R-phycoerythrin (PE) (Molecular Probes, Eugene, OR), FITC, Rhodamine, and Texas Red (Pierce, Rockford, IL), Cy5, Cy5.5, Cy7 (Amersham Life Science, Pittsburgh, PA). Suitable optical dyes, including fluorophores, are described in Molecular Probes Handbook by Richard P. Haugland, hereby expressly incorporated by reference.

Suitable proteinaceous fluorescent labels also include, but are not limited to, green fluorescent protein, including a Renilla, Ptilosarcus, or Aequorea species of GFP (Chalfie *et al.*, 1994, *Science* 263:802-805), EGFP (Clontech Laboratories, Inc., Genbank Accession Number U55762), blue fluorescent protein (BFP, Quantum Biotechnologies, Inc. 1801 de Maisonneuve Blvd. West, 8th Floor, Montreal, Quebec, Canada H3H 1J9; Stauber, 1998, *Biotechniques* 24:462-471; Heim *et al.*, 1996, *Curr. Biol.* 6:178-182), enhanced yellow fluorescent protein (EYFP, Clontech Laboratories, Inc.), luciferase (Ichiki *et al.*, 1993, *J. Immunol.* 150:5408-5417), β galactosidase (Nolan *et al.*, 1988, *Proc. Natl. Acad. Sci. U.S.A.* 85:2603-2607) and Renilla (WO92/15673, WO95/07463, WO98/14605, WO98/26277, WO99/49019, U.S. Patent Nos. 5292658, 5418155, 5683888, 5741668, 5777079, 5804387, 5874304, 5876995, 5925558). All of the above-cited references are expressly incorporated herein by reference.

A CD33 targeting compound described herein may also comprise additional domains, which e.g. are helpful in the isolation of the molecule or relate to an adapted pharmacokinetic profile of the molecule.

Domains helpful for the isolation of an antibody construct may be elected from peptide motives or secondarily introduced moieties, which can be captured in an isolation method, e.g. an isolation column. A non-limiting embodiments of such additional domains comprise peptide motives known as Myc-tag, HAT-tag, HA-tag, TAP-tag, GST-tag, chitin binding domain (CBD-tag), maltose binding protein (MBP-tag), Flag-tag, Strep-tag and variants thereof (e.g. StreptII-tag) and His-tag. All herein disclosed antibody constructs characterized by the identified CDRs are preferred to comprise a His-tag domain, which is generally known as a repeat of consecutive His residues in the amino acid sequence of a molecule, preferably of six His residues.

The term "epigenetic factor" in connection with the present invention defines a compound which is capable of changing the gene expression or cellular phenotype of a cell population upon administration. It is understood that such change refers to one or more functional relevant modifications to the genome without involving a change in the nucleic acid sequence. Examples of such modifications are DNA methylation and histone modification, which are both important for the regulation of gene expression without altering the underlying DNA sequence. Particular examples for epigenetic factors suitable in the combination therapy approach according to the invention are selected from the group consisting of histone deacetylase (HDAC) inhibitors, DNA methyltransferase (DNMT) I inhibitors, hydroxyurea, Granulocyte-Colony Stimulating Factor (G-CSF), histone demethylase inhibitors and ATRA (All Trans-retinoic acid).

Histone deacetylases (HDACs) are a class of enzymes, that remove acetyl groups from ϵ -N-acetyl lysine amino acid on a histone. The deacetylated histone is suitable to wrap the DNA more tightly which negatively effects the expression genes in the region of the wrapped DNA. Accordingly, HDAC inhibitors inhibit the enzymatic deacetylation of histone and allow for the expression or enhancement of expression of those genes located in the region of the acetylated histone. Non-limiting examples for HDAC inhibitors in connection with this invention comprise panobinostat, vorinostat, romidepsin, N-acetyldinaline, belinostat, givinostat, entinostat, mocetinostat, EVP-0334, SRT501, CUDC-101, Quisinostat, abexinostat, LAQ824, and valproic acid.

DNA methyltransferase (DNMT) I is an enzyme, which catalyze the transfer of a methyl group to DNA. The degree of DNA methylation is also decisive for the expression of genes. Non-limiting examples for DNMT I inhibitors in connection with this invention comprise 5-azacitidine, decitabine, hydralazine, zebularine, procainamide, (-)-epigallocatechin-3-gallate, MG98, RG108, and SGI-110.

Histone methyltransferases (HMT) are histone-modifying enzymes that catalyze the transfer of one, two, or three methyl groups to lysine and arginine residues of histone proteins. Methylation of histones is important biologically because it is the principal epigenetic modification of chromatin that determines gene expression, genomic stability, stem cell maturation, cell lineage development, genetic imprinting, DNA methylation, and cell mitosis. Non-limiting examples for HMT inhibitors in connection with this invention comprise LSD1 (KDM1A) demethylase inhibitor, and chaetocin.

ATRA (All Trans-retinoic acid) is the carboxylic acid form of vitamin A and is also known as Tretinoin.

Hydroxyurea is also known as Hydroxycarbamide and is used as an antineoplastic drug. The compound is described to decrease the production of deoxyribonucleotides via inhibition of the enzyme ribonucleotide reductase.

Granulocyte-colony stimulating factor (G-CSF or GCSF), also known as colony-stimulating factor 3 (CSF 3), is a glycoprotein that stimulates the bone marrow to produce granulocytes and stem cells and release them into the bloodstream. Functionally, it is a cytokine and hormone, a type of colony-stimulating factor, and is produced by a number of different tissues. The pharmaceutical analogs of naturally occurring G-CSF are called filgrastim and lenograstim.

The term "nucleic acid" is well known to the skilled person and encompasses DNA (such as cDNA) and RNA (such as mRNA). The nucleic acid can be double stranded and single stranded, linear and circular. Said nucleic acid molecule is preferably comprised in a vector which is preferably comprised in a host cell. Said host cell is, e.g. after transformation or transfection with the nucleic acid sequence described herein, capable of expressing the CD33 targeting compound. For that purpose the nucleic acid molecule is operatively linked with control sequences.

A vector is a nucleic acid molecule used as a vehicle to transfer (foreign) genetic material into a cell. The term "vector" encompasses – but is not restricted to – plasmids, viruses, cosmids and artificial chromosomes. In general, engineered vectors comprise an origin of replication, a multicloning site and a selectable marker. The vector itself is generally a nucleotide sequence, commonly a DNA sequence that comprises an insert (transgene) and a larger sequence that serves as the "backbone" of the vector. Modern vectors may encompass additional features besides the transgene insert and a backbone: promoter, genetic marker, antibiotic resistance, reporter gene, targeting sequence, protein purification tag. Vectors called expression vectors (expression constructs) specifically are for the expression of the transgene in the target cell, and generally have control sequences such as a promoter sequence that drives expression of the transgene. Insertion of a vector into the target cell is usually called "transformation" for bacteria, "transfection" for eukaryotic cells, although insertion of a viral vector is also called "transduction".

As used herein, the term "host cell" is intended to refer to a cell into which a nucleic acid encoding the CD33 targeting compound described herein is introduced by way of transformation, transfection and the like. It should be understood that such terms refer not only to the particular subject cell but to the progeny or potential progeny of such a cell. Because certain modifications may occur in succeeding generations due to either mutation

or environmental influences, such progeny may not, in fact, be identical to the parent cell, but are still included within the scope of the term as used herein.

As used herein, the term "expression" includes any step involved in the production of a CD33 targeting compound described herein including, but not limited to, transcription, post-transcriptional modification, translation, post-translational modification, and secretion.

The term "control sequences" refers to DNA sequences necessary for the expression of an operably linked coding sequence in a particular host organism. The control sequences that are suitable for prokaryotes, for example, include a promoter, optionally an operator sequence, and a ribosome binding site. Eukaryotic cells are known to utilize promoters, polyadenylation signals, and enhancers.

A nucleic acid is "operably linked" when it is placed into a functional relationship with another nucleic acid sequence. For example, DNA for a presequence or secretory leader is operably linked to DNA for a polypeptide if it is expressed as a preprotein that participates in the secretion of the polypeptide; a promoter or enhancer is operably linked to a coding sequence if it affects the transcription of the sequence; or a ribosome binding site is operably linked to a coding sequence if it is positioned so as to facilitate translation. Generally, "operably linked" means that the DNA sequences being linked are contiguous, and, in the case of a secretory leader, contiguous and in reading phase. However, enhancers do not have to be contiguous. Linking is accomplished by ligation at convenient restriction sites. If such sites do not exist, the synthetic oligonucleotide adaptors or linkers are used in accordance with conventional practice.

The terms "host cell," "target cell" or "recipient cell" are intended to include any individual cell or cell culture that can be or has/have been recipients for vectors or the incorporation of exogenous nucleic acid molecules, polynucleotides and/or proteins. It also is intended to include progeny of a single cell, and the progeny may not necessarily be completely identical (in morphology or in genomic or total DNA complement) to the original parent cell due to natural, accidental, or deliberate mutation. The cells may be prokaryotic or eukaryotic, and include but are not limited to bacteria, yeast cells, animal cells, and mammalian cells, e.g., murine, rat, macaque or human.

Suitable eukaryotic host cells include yeasts, fungi, insect cells and mammalian cells.

The CD33 targeting compound described herein can be produced in bacteria. After expression, the CD33 targeting compound, preferably the antibody construct is isolated from the *E. coli* cell paste in a soluble fraction and can be purified through, e.g., affinity chromatography and/or size exclusion. Final purification can be carried out similar to the process for purifying antibody expressed e. g., in CHO cells.

In addition to prokaryotes, eukaryotic microbes such as filamentous fungi or yeast are suitable cloning or expression hosts for the CD33 targeting compound described herein. *Saccharomyces cerevisiae*, or common baker's yeast, is the most commonly used among lower eukaryotic host microorganisms. However, a number of other genera, species, and strains are commonly available and useful herein, such as *Schizosaccharomyces pombe*, Kluyveromyces hosts such as, e.g., *K. lactis*, *K. fragilis* (ATCC 12424), *K. bulgaricus* (ATCC 16045), *K. wickerhamii* (ATCC 24178), *K. waltii* (ATCC 56500), *K. drosophilarum* (ATCC 36906), *K. thermotolerans*, and *K. marxianus*; yarrowia (EP 402 226); *Pichia pastoris* (EP 183 070); *Candida*; *Trichoderma reesia* (EP 244 234); *Neurospora crassa*; Schwanniomyces such as *Schwanniomyces occidentalis*; and filamentous fungi such as, e.g., *Neurospora*, *Penicillium*, *Tolypocladium*, and *Aspergillus* hosts such as *A. nidulans* and *A. niger*.

Suitable host cells for the expression of glycosylated CD33 targeting compounds described herein, preferably antibody derived antibody constructs are derived from multicellular organisms. Examples of invertebrate cells include plant and insect cells. Numerous baculoviral strains and variants and corresponding permissive insect host cells from hosts such as *Spodoptera frugiperda* (caterpillar), *Aedes aegypti* (mosquito), *Aedes albopictus* (mosquito), *Drosophila melanogaster* (fruit fly), and *Bombyx mori* have been identified. A variety of viral strains for transfection are publicly available, e. g. , the L-1 variant of *Autographa californica* NPV and the Bm-5 strain of *Bombyx mori* NPV, and such viruses may be used as the virus herein according to the present invention, particularly for transfection of *Spodoptera frugiperda* cells.

Plant cell cultures of cotton, corn, potato, soybean, petunia, tomato, Arabidopsis and tobacco can also be utilized as hosts. Cloning and expression vectors useful in the production of proteins in plant cell culture are known to those of skill in the art. See e.g. Hiatt *et al.*, Nature (1989) 342: 76-78, Owen *et al.* (1992) Bio/Technology 10: 790-794, Artsaenko *et al.* (1995) The Plant J 8: 745-750, and Fecker *et al.* (1996) Plant Mol Biol 32: 979-986.

However, interest has been greatest in vertebrate cells, and propagation of vertebrate cells in culture (tissue culture) has become a routine procedure. Examples of useful mammalian host cell lines are monkey kidney CV1 line transformed by SV40 (COS-7, ATCC CRL 1651); human embryonic kidney line (293 or 293 cells subcloned for growth in suspension culture, Graham *et al.* , J. Gen Virol. 36 : 59 (1977)); baby hamster kidney cells (BHK, ATCC CCL 10); Chinese hamster ovary cells/-DHFR (CHO, Urlaub *et al.* , Proc. Natl. Acad. Sci. USA 77: 4216 (1980)); mouse sertoli cells (TM4, Mather, Biol. Reprod. 23: 243-251 (1980)); monkey kidney cells (CV1 ATCC CCL 70); African green monkey kidney cells (VERO-76, ATCC CRL1587) ; human cervical carcinoma cells (HELA, ATCC CCL 2); canine kidney cells (MDCK, ATCC CCL 34); buffalo rat liver cells (BRL 3A, ATCC CRL 1442); human lung cells

(W138, ATCC CCL 75); human liver cells (Hep G2, 1413 8065); mouse mammary tumor (MMT 060562, ATCC CCL5 1); TRI cells (Mather *et al.*, Annals N. Y Acad. Sci. 383 : 44-68 (1982)) ; MRC 5 cells; FS4 cells; and a human hepatoma line (Hep G2).

When using recombinant techniques, the CD33 targeting compound described herein can be produced intracellularly, in the periplasmic space, or directly secreted into the medium. If the antibody construct is produced intracellularly, as a first step, the particulate debris, either host cells or lysed fragments, are removed, for example, by centrifugation or ultrafiltration. Carter *et al.*, Bio/Technology 10: 163-167 (1992) describe a procedure for isolating antibodies which are secreted to the periplasmic space of *E. coli*. Briefly, cell paste is thawed in the presence of sodium acetate (pH 3.5), EDTA, and phenylmethylsulfonylfluoride (PMSF) over about 30 min. Cell debris can be removed by centrifugation. Where the antibody is secreted into the medium, supernatants from such expression systems are generally first concentrated using a commercially available protein concentration filter, for example, an Amicon or Millipore Pellicon ultrafiltration unit. A protease inhibitor such as PMSF may be included in any of the foregoing steps to inhibit proteolysis and antibiotics may be included to prevent the growth of adventitious contaminants.

The CD33 targeting compound described herein prepared from the host cells can be purified using, for example, hydroxylapatite chromatography, gel electrophoresis, dialysis, and affinity chromatography, with affinity chromatography being the preferred purification technique.

The matrix to which the affinity ligand is attached is most often agarose, but other matrices are available. Mechanically stable matrices such as controlled pore glass or poly(styrene-divinyl) benzene allow for faster flow rates and shorter processing times than can be achieved with agarose. Where the CD33 targeting compound described herein comprises a CH3 domain, the Bakerbond ABXMresin (J. T. Baker, Phillipsburg, NJ) is useful for purification. Other techniques for protein purification such as fractionation on an ion-exchange column, ethanol precipitation, Reverse Phase HPLC, chromatography on silica, chromatography on heparin SEPHAROSE™ chromatography on an anion or cation exchange resin (such as a polyaspartic acid column), chromato-focusing, SDS-PAGE, and ammonium sulfate precipitation are also available depending on the antibody to be recovered.

The term "culturing" refers to the in vitro maintenance, differentiation, growth, proliferation and/or propagation of cells under suitable conditions in a medium.

As used herein, the term "pharmaceutical composition" relates to a composition for administration to a patient, preferably a human patient. The particular preferred

pharmaceutical composition of this invention comprises a CD33 targeting compound and at least one epigenetic factor, either in one single formulation or in separate formulations. Preferably, the pharmaceutical composition comprises suitable formulations of carriers, stabilizers and/or excipients. In a preferred embodiment, the pharmaceutical composition comprises a composition for parenteral, transdermal, intraluminal, intraarterial, intrathecal and/or intranasal administration or by direct injection into tissue. It is in particular envisaged that said composition is administered to a patient via infusion or injection. Administration of the suitable compositions may be effected by different ways, e.g., by intravenous, intraperitoneal, subcutaneous, intramuscular, topical or intradermal administration. In particular, the present invention provides for an uninterrupted administration of the suitable composition. As a non-limiting example, uninterrupted, i.e. continuous administration may be realized by a small pump system worn by the patient for metering the influx of therapeutic agent into the body of the patient. The pharmaceutical composition comprising the CD33 targeting compound or the CD33 targeting compound and at least one epigenetic factor can be administered by using said pump systems. Such pump systems are generally known in the art, and commonly rely on periodic exchange of cartridges containing the therapeutic agent to be infused. When exchanging the cartridge in such a pump system, a temporary interruption of the otherwise uninterrupted flow of therapeutic agent into the body of the patient may ensue. In such a case, the phase of administration prior to cartridge replacement and the phase of administration following cartridge replacement would still be considered within the meaning of the pharmaceutical means and methods of the invention together make up one "uninterrupted administration" of such therapeutic agent.

The continuous or uninterrupted administration of the CD33 targeting compound described herein or the CD33 targeting compound and at least one epigenetic factor described herein may be intravenous or subcutaneous by way of a fluid delivery device or small pump system including a fluid driving mechanism for driving fluid out of a reservoir and an actuating mechanism for actuating the driving mechanism. Pump systems for subcutaneous administration may include a needle or a cannula for penetrating the skin of a patient and delivering the suitable composition into the patient's body. Said pump systems may be directly fixed or attached to the skin of the patient independently of a vein, artery or blood vessel, thereby allowing a direct contact between the pump system and the skin of the patient. The pump system can be attached to the skin of the patient for 24 hours up to several days. The pump system may be of small size with a reservoir for small volumes. As a non-limiting example, the volume of the reservoir for the suitable pharmaceutical composition to be administered can be between 0.1 and 50 ml.

The continuous administration may be transdermal by way of a patch worn on the skin and replaced at intervals. One of skill in the art is aware of patch systems for drug delivery suitable for this purpose. It is of note that transdermal administration is especially amenable to uninterrupted administration, as exchange of a first exhausted patch can advantageously be accomplished simultaneously with the placement of a new, second patch, for example on the surface of the skin immediately adjacent to the first exhausted patch and immediately prior to removal of the first exhausted patch. Issues of flow interruption or power cell failure do not arise.

The inventive compositions may further comprise a pharmaceutically acceptable carrier.

Examples of suitable pharmaceutical carriers are well known in the art and include solutions, e.g. phosphate buffered saline solutions, water, emulsions, such as oil/water emulsions, various types of wetting agents, sterile solutions, liposomes, etc. Compositions comprising such carriers can be formulated by well known conventional methods. Formulations can comprise carbohydrates, buffer solutions, amino acids and/or surfactants. Carbohydrates may be non-reducing sugars, preferably trehalose, sucrose, octasulfate, sorbitol or xylitol. In general, as used herein, "pharmaceutically acceptable carrier" means any and all solvents, dispersion media, coatings, antibacterial and antifungal agents, isotonic and absorption delaying agents, compatible with pharmaceutical administration. The use of such media and agents for pharmaceutically active substances is well known in the art. Acceptable carriers, excipients, or stabilizers are nontoxic to recipients at the dosages and concentrations employed and include: additional buffering agents; preservatives; co-solvents; antioxidants, including ascorbic acid and methionine; chelating agents such as EDTA; metal complexes (e.g., Zn-protein complexes); biodegradable polymers, such as polyesters; salt-forming counter-ions, such as sodium, polyhydric sugar alcohols; amino acids, such as alanine, glycine, asparagine, 2-phenylalanine, and threonine; sugars or sugar alcohols, such as trehalose, sucrose, octasulfate, sorbitol or xylitol, stachyose, mannose, sorbose, xylose, ribose, myoinositol, galactose, lactitol, ribitol, myoinositol, galactitol, glycerol, cyclitols (e.g., inositol), polyethylene glycol; sulfur containing reducing agents, such as glutathione, thiocetic acid, sodium thioglycolate, thioglycerol, [alpha]-monothioglycerol, and sodium thio sulfate; low molecular weight proteins, such as human serum albumin, bovine serum albumin, gelatin, or other immunoglobulins; and hydrophilic polymers, such as polyvinylpyrrolidone. Such formulations may be used for continuous administrations which may be intravenous or subcutaneous with and/or without pump systems. Amino acids may be charged amino acids, preferably lysine, lysine acetate, arginine, glutamate and/or histidine. Surfactants may be detergents, preferably with a molecular weight of >1.2 KD and/or a polyether, preferably with a molecular weight of >3 KD. Non-limiting examples for preferred detergents are Tween

20, Tween 40, Tween 60, Tween 80 or Tween 85. Non-limiting examples for preferred polyethers are PEG 3000, PEG 3350, PEG 4000 or PEG 5000. Buffer systems used in the present invention can have a preferred pH of 5-9 and may comprise citrate, succinate, phosphate, histidine and acetate.

5 The compositions of the present invention comprising the CD33 targeting compound and at least one epigenetic factor in a single or separate formulations can be administered to the subject at a suitable dose which can be determined e.g. by dose escalating studies by administration of increasing doses of the polypeptide described herein exhibiting cross-species specificity described herein to non-chimpanzee primates, for instance macaques. As
10 set forth above, the CD33 targeting composition described herein exhibiting cross-species specificity described herein can be advantageously used in identical form in preclinical testing in non-chimpanzee primates and as drug in humans. The composition or these compositions can also be administered in combination with additional other proteinaceous and non-proteinaceous drugs. These drugs may be administered simultaneously with the
15 composition comprising the polypeptide described herein as defined herein or separately before or after administration of said polypeptide in timely defined intervals and doses. The dosage regimen will be determined by the attending physician and clinical factors. As is well known in the medical arts, dosages for any one patient depend upon many factors, including the patient's size, body surface area, age, the particular compound to be administered, sex,
20 time and route of administration, general health, and other drugs being administered concurrently.

Preparations for parenteral administration include sterile aqueous or non-aqueous solutions, suspensions, and emulsions. Examples of non-aqueous solvents are propylene glycol, polyethylene glycol, vegetable oils such as olive oil, and injectable organic esters such as
25 ethyl oleate. Aqueous carriers include water, alcoholic/aqueous solutions, emulsions or suspensions, including saline and buffered media. Parenteral vehicles include sodium chloride solution, Ringer's dextrose, dextrose and sodium chloride, lactated Ringer's, or fixed oils. Intravenous vehicles include fluid and nutrient replenishers, electrolyte replenishers (such as those based on Ringer's dextrose), and the like. Preservatives and other additives
30 may also be present such as, for example, antimicrobials, anti-oxidants, chelating agents, inert gases and the like. In addition, the composition of the present invention might comprise proteinaceous carriers, like, e.g., serum albumin or immunoglobulin, preferably of human origin. It is envisaged that the composition of the invention might comprise, in addition to the polypeptide described herein defined herein, further biologically active agents, depending on
35 the intended use of the composition. Such agents might be drugs acting on the gastrointestinal system, drugs acting as cytostatica, drugs preventing hyperuricemia, drugs

inhibiting immunoreactions (e.g. corticosteroids), drugs modulating the inflammatory response, drugs acting on the circulatory system and/or agents such as cytokines known in the art. It is also envisaged that the composition of the present invention comprising the CD33 targeting compound and at least one epigenetic factor in a single or separate formulations is applied in an additional co-therapy, i.e., in combination with another anti-cancer medicament.

The biological activity of the pharmaceutical composition defined herein can be determined for instance by cytotoxicity assays, as described in the following examples, in WO 99/54440 or by Schlereth *et al.* (Cancer Immunol. Immunother. 20 (2005), 1-12). "Efficacy" or "*in vivo* efficacy" as used herein refers to the response to therapy by the pharmaceutical composition of the invention, using e.g. standardized NCI response criteria. The success or *in vivo* efficacy of the therapy using a pharmaceutical composition of the invention refers to the effectiveness of the composition for its intended purpose, i.e. the ability of the composition to cause its desired effect, i.e. depletion of pathologic cells, e.g. tumor cells. The *in vivo* efficacy may be monitored by established standard methods for the respective disease entities including, but not limited to white blood cell counts, differentials, Fluorescence Activated Cell Sorting, bone marrow aspiration. In addition, various disease specific clinical chemistry parameters and other established standard methods may be used. Furthermore, computer-aided tomography, X-ray, nuclear magnetic resonance tomography (e.g. for National Cancer Institute-criteria based response assessment [Cheson BD, Horning SJ, Coiffier B, Shipp MA, Fisher RI, Connors JM, Lister TA, Vose J, Grillo-Lopez A, Hagenbeek A, Cabanillas F, Klippensten D, Hiddemann W, Castellino R, Harris NL, Armitage JO, Carter W, Hoppe R, Canellos GP. Report of an international workshop to standardize response criteria for non-Hodgkin's lymphomas. NCI Sponsored International Working Group. J Clin Oncol. 1999 Apr;17(4):1244]), positron-emission tomography scanning, white blood cell counts, differentials, Fluorescence Activated Cell Sorting, bone marrow aspiration, lymph node biopsies/histologies, and various lymphoma specific clinical chemistry parameters (e.g. lactate dehydrogenase) and other established standard methods may be used.

Another major challenge in the development of drugs such as the pharmaceutical composition of the invention is the predictable modulation of pharmacokinetic properties. To this end, a pharmacokinetic profile of the drug candidate, i.e. a profile of the pharmacokinetic parameters that affect the ability of a particular drug to treat a given condition, can be established. Pharmacokinetic parameters of the drug influencing the ability of a drug for treating a certain disease entity include, but are not limited to: half-life, volume of distribution, hepatic first-pass metabolism and the degree of blood serum binding. The efficacy of a given drug agent can be influenced by each of the parameters mentioned above.

"Half-life" means the time where 50% of an administered drug are eliminated through biological processes, e.g. metabolism, excretion, etc.

By "hepatic first-pass metabolism" is meant the propensity of a drug to be metabolized upon first contact with the liver, i.e. during its first pass through the liver.

- 5 "Volume of distribution" means the degree of retention of a drug throughout the various compartments of the body, like e.g. intracellular and extracellular spaces, tissues and organs, etc. and the distribution of the drug within these compartments.

10 "Degree of blood serum binding" means the propensity of a drug to interact with and bind to blood serum proteins, such as albumin, leading to a reduction or loss of biological activity of the drug.

Pharmacokinetic parameters also include bioavailability, lag time (T_{lag}), T_{max}, absorption rates, more onset and/or C_{max} for a given amount of drug administered. "Bioavailability" means the amount of a drug in the blood compartment. "Lag time" means the time delay between the administration of the drug and its detection and measurability in blood or plasma.

15 "T_{max}" is the time after which maximal blood concentration of the drug is reached, and "C_{max}" is the blood concentration maximally obtained with a given drug. The time to reach a blood or tissue concentration of the drug which is required for its biological effect is influenced by all parameters. Pharmacokinetic parameters of comprising a CD33 targeting compound such as a bispecific single chain antibody exhibiting cross-species specificity, which may be determined in preclinical animal testing in non-chimpanzee primates as outlined above, are also set forth e.g. in the publication by Schlereth *et al.* (Cancer Immunol. Immunother. 20 (2005), 1-12).

25 The term "toxicity" as used herein refers to the toxic effects of a drug manifested in adverse events or severe adverse events. These side events might refer to a lack of tolerability of the drug in general and/or a lack of local tolerance after administration. Toxicity could also include teratogenic or carcinogenic effects caused by the drug.

30 The term "safety", "*in vivo* safety" or "tolerability" as used herein defines the administration of a drug without inducing severe adverse events directly after administration (local tolerance) and during a longer period of application of the drug. "Safety", "*in vivo* safety" or "tolerability" can be evaluated e.g. at regular intervals during the treatment and follow-up period. Measurements include clinical evaluation, e.g. organ manifestations, and screening of laboratory abnormalities. Clinical evaluation may be carried out and deviations to normal findings recorded/coded according to NCI-CTC and/or MedDRA standards. Organ

manifestations may include criteria such as allergy/immunology, blood/bone marrow, cardiac arrhythmia, coagulation and the like, as set forth e.g. in the Common Terminology Criteria for adverse events v3.0 (CTCAE). Laboratory parameters which may be tested include for instance hematology, clinical chemistry, coagulation profile and urine analysis and examination of other body fluids such as serum, plasma, lymphoid or spinal fluid, liquor and the like. Safety can thus be assessed e.g. by physical examination, imaging techniques (i.e. ultrasound, x-ray, CT scans, Magnetic Resonance Imaging (MRI), other measures with technical devices (i.e. electrocardiogram), vital signs, by measuring laboratory parameters and recording adverse events. For example, adverse events in non-chimpanzee primates in the uses and methods according to the invention may be examined by histopathological and/or histochemical methods.

The term "effective dose" or "effective dosage" is defined as an amount sufficient to achieve or at least partially achieve the desired effect. The term "therapeutically effective dose" is defined as an amount sufficient to cure or at least partially arrest the disease and its complications in a patient already suffering from the disease. Amounts effective for this use will depend upon the severity of the infection and the general state of the subject's own immune system. The term "patient" includes human and other mammalian subjects that receive either prophylactic or therapeutic treatment.

The term "effective and non-toxic dose" as used herein refers to a tolerable dose of a pharmaceutical composition (i.e. a pharmaceutical composition comprising the CD33 targeting compound and at least one epigenetic factor in a single or separate formulations) which is high enough to cause depletion of pathologic cells, tumor elimination, tumor shrinkage or stabilization of disease without or essentially without major toxic effects. Such effective and non-toxic doses may be determined e.g. by dose escalation studies described in the art and should be below the dose inducing severe adverse side events (dose limiting toxicity, DLT).

The above terms are also referred to e.g. in the Preclinical safety evaluation of biotechnology-derived pharmaceuticals S6; ICH Harmonised Tripartite Guideline; ICH Steering Committee meeting on July 16, 1997.

The appropriate dosage, or therapeutically effective amount, of a pharmaceutical composition comprising the CD33 targeting compound and at least one epigenetic factor in a single or separate formulations will depend on the condition to be treated, the severity of the condition, prior therapy, and the patient's clinical history and response to the therapeutic agent. The proper dose can be adjusted according to the judgment of the attending physician such that it can be administered to the patient one time or over a series of

administrations. The pharmaceutical composition can be administered as a sole therapeutic or in combination with additional therapies such as anti-cancer therapies as needed.

The pharmaceutical compositions of this invention are particularly useful for parenteral administration, i.e., subcutaneously, intramuscularly, intravenously, intra-articular and/or intra-synovial. Parenteral administration can be by bolus injection or continuous infusion.

If the pharmaceutical composition has been lyophilized, the lyophilized material is first reconstituted in an appropriate liquid prior to administration. The lyophilized material may be reconstituted in, e.g., bacteriostatic water for injection (BWI), physiological saline, phosphate buffered saline (PBS), or the same formulation the protein had been in prior to lyophilization.

It has been surprisingly found in connection with the present invention that specific groups of epigenetic factors relate to an increase of the potency of a therapy approach making use of the engagement of T cells to CD33 positive target cells. 5-azacytidine (Vidaza™) and 5-aza-2'-deoxycytidine (decitabine, Dacogen™) are nucleoside analogs that belong to a class of epigenetic therapeutics and are known to be capable of inducing tumor cell killing through the disruption of protein synthesis and inhibition of DNA methylation. Co-therapy approaches using the CD33 specific antibody lintuzumab and 5-azacytidine lead to an improved lintuzumab-mediated ADCC against AML target cells. However, in such study the incubation of AML cells or macrophages did not affect the CD33 expression on those cells (Sutherland et al. MAbs. 2010 Jul-Aug; 2(4): 440–448). Accordingly it has been concluded that since for the mode-of-action of lintuzumab requires a functional interaction between the F_C domain of lintuzumab and the F_Cγ receptor of immune effector cells is required the pretreatment with epigenetic compounds increases the specific phagocytosis of target cells by the effector cells.

Thus the present finding are especially in the light of those known effect surprisingly, since the mode-of-action underlying the target cells lysis by T cells when engaged by the CD33 targeting compounds described herein is completely independent of any F_C-F_Cγ receptor interaction. In contrast to the previous studies, an increase of CD33 surface expression by treatment of target cells with epigenetic factors described herein was observed. Also, as apparent from the mode-of-action underlying the target cells lysis by T cells the observed synergistic effect is independent of any signal-transduction-effect which might be triggered by antibody binding to the CD33 surface molecule (e.g. a signaling pathway involving SHP-1, Syk, or both).

Moreover, as the speed and extent of myeloid leukemia cell blast lysis at a given time point was proportional to CD33 expression, this finding supports that the combination of an

epigenetic therapy and a CD33 directed T cell engager would be synergistically more effective than either therapy administered separately. Accordingly, the described administration of one or more CD33 targeting compound in combination with epigenetic factors described herein may allow for lower doses of a bispecific T cell engager to be effective at a given time point.

As described above, examples for epigenetic factors suitable in the combination therapy approach according to the invention are selected from the group consisting of histone deacetylase (HDAC) inhibitors, DNA methyltransferase (DNMT) I inhibitors, hydroxyurea, Granulocyte-Colony Stimulating Factor (G-CSF), histone demethylase inhibitors and ATRA (All Trans-retinoic acid). The redirected lysis of target cells via the recruitment of T cells by a multispecific, at least bispecific, construct involves cytolytic synapse formation and delivery of perforin and granzymes. The engaged T cells are capable of serial target cell lysis, and are not affected by immune escape mechanisms interfering with peptide antigen processing and presentation, or clonal T cell differentiation; see, for example, WO 2007/042261.

The cytotoxic activity mediated by CD33/CD3 bispecific compounds described herein such as bispecific antibody constructs is preferably measured in a cell-based cytotoxicity assay. It is represented by the EC_{50} value, which corresponds to the half maximal effective concentration (concentration of the compound which induces a cytotoxic response halfway between the baseline and maximum). Preferably, the EC_{50} value of the CD33/CD3 bispecific antibody constructs is $\leq 20,000$ pg/ml, more preferably ≤ 5000 pg/ml, even more preferably ≤ 1000 pg/ml, even more preferably ≤ 500 pg/ml, even more preferably ≤ 350 pg/ml, even more preferably ≤ 320 pg/ml, even more preferably ≤ 250 pg/ml, even more preferably ≤ 100 pg/ml, even more preferably ≤ 50 pg/ml, even more preferably ≤ 10 pg/ml, and most preferably ≤ 5 pg/ml.

Any of the above given EC_{50} values can be combined with any one of the indicated scenarios of a cell-based cytotoxicity assay. For example, when (human) CD8 positive T cells or a macaque T cell line are used as effector cells, the EC_{50} value of the CD33/CD3 bispecific antibody construct is preferably ≤ 1000 pg/ml, more preferably ≤ 500 pg/ml, even more preferably ≤ 250 pg/ml, even more preferably ≤ 100 pg/ml, even more preferably ≤ 50 pg/ml, even more preferably ≤ 10 pg/ml, and most preferably ≤ 5 pg/ml. If in this assay the target cells are (human or macaque) CD33 transfected cells such as CHO cells, the EC_{50} value of the CD33/CD3 bispecific antibody construct is preferably ≤ 150 pg/ml, more preferably ≤ 100 pg/ml, even more preferably ≤ 50 pg/ml, even more preferably ≤ 30 pg/ml, even more preferably ≤ 10 pg/ml, and most preferably ≤ 5 pg/ml.

If the target cells are a CD33 positive natural expresser cell line, then the EC₅₀ value is preferably ≤350 pg/ml, more preferably ≤320 pg/ml, even more preferably ≤250 pg/ml, even more preferably ≤200 pg/ml, even more preferably ≤100 pg/ml, even more preferably ≤150 pg/ml, even more preferably ≤100 pg/ml, and most preferably ≤50 pg/ml, or lower.

- 5 When (human) PBMCs are used as effector cells, the EC₅₀ value of the CD33/CD3 bispecific antibody construct is preferably ≤1000 pg/ml, more preferably ≤750 pg/ml, more preferably ≤500 pg/ml, even more preferably ≤350 pg/ml, even more preferably ≤320 pg/ml, even more preferably ≤250 pg/ml, even more preferably ≤100 pg/ml, and most preferably ≤50 pg/ml, or lower.
- 10 The difference in cytotoxic activity between the monomeric and the dimeric isoform of individual CD33/CD3 bispecific antibody constructs is referred to as "potency gap". This potency gap can e.g. be calculated as ratio between EC₅₀ values of the molecule's monomeric and dimeric form. Potency gaps of the CD33/CD3 bispecific antibody constructs described herein are preferably ≤5, more preferably ≤4, even more preferably ≤3, even more
- 15 preferably ≤2 and most preferably ≤1.

It is particularly preferred for the CD33 targeting compound described herein that the second binding domain capable of binding to the T cell CD3 receptor complex comprises a VL region comprising CDR-L1, CDR-L2 and CDR-L3 selected from:

- (a) CDR-L1 as depicted in SEQ ID NO: 27 of WO 2008/119567, CDR-L2 as depicted in
- 20 SEQ ID NO: 28 of WO 2008/119567 and CDR-L3 as depicted in SEQ ID NO: 29 of WO 2008/119567;
- (b) CDR-L1 as depicted in SEQ ID NO: 117 of WO 2008/119567, CDR-L2 as depicted in SEQ ID NO: 118 of WO 2008/119567 and CDR-L3 as depicted in SEQ ID NO: 119 of WO 2008/119567; and
- 25 (c) CDR-L1 as depicted in SEQ ID NO: 153 of WO 2008/119567, CDR-L2 as depicted in SEQ ID NO: 154 of WO 2008/119567 and CDR-L3 as depicted in SEQ ID NO: 155 of WO 2008/119567.

- In an alternatively preferred embodiment of the CD33 targeting compound described herein, the second binding domain capable of binding to the T cell CD3 receptor complex comprises
- 30 a VH region comprising CDR-H 1, CDR-H2 and CDR-H3 selected from:

- (a) CDR-H1 as depicted in SEQ ID NO: 12 of WO 2008/119567, CDR-H2 as depicted in SEQ ID NO: 13 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 14 of WO 2008/119567;
- (b) CDR-H1 as depicted in SEQ ID NO: 30 of WO 2008/119567, CDR-H2 as depicted in
- 35 SEQ ID NO: 31 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 32 of WO 2008/119567;

(c) CDR-H1 as depicted in SEQ ID NO: 48 of WO 2008/119567, CDR-H2 as depicted in SEQ ID NO: 49 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 50 of WO 2008/119567;

(d) CDR-H1 as depicted in SEQ ID NO: 66 of WO 2008/119567, CDR-H2 as depicted in SEQ ID NO: 67 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 68 of WO 2008/119567;

(e) CDR-H1 as depicted in SEQ ID NO: 84 of WO 2008/119567, CDR-H2 as depicted in SEQ ID NO: 85 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 86 of WO 2008/119567;

(f) CDR-H1 as depicted in SEQ ID NO: 102 of WO 2008/119567, CDR-H2 as depicted in SEQ ID NO: 103 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 104 of WO 2008/119567;

(g) CDR-H1 as depicted in SEQ ID NO: 120 of WO 2008/119567, CDR-H2 as depicted in SEQ ID NO: 121 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 122 of WO 2008/119567;

(h) CDR-H1 as depicted in SEQ ID NO: 138 of WO 2008/119567, CDR-H2 as depicted in SEQ ID NO: 139 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 140 of WO 2008/119567;

(i) CDR-H1 as depicted in SEQ ID NO: 156 of WO 2008/119567, CDR-H2 as depicted in SEQ ID NO: 157 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 158 of WO 2008/119567; and

(j) CDR-H1 as depicted in SEQ ID NO: 174 of WO 2008/119567, CDR-H2 as depicted in SEQ ID NO: 175 of WO 2008/119567 and CDR-H3 as depicted in SEQ ID NO: 176 of WO 2008/119567.

It is further preferred for the CD33 targeting compound described herein that the second binding domain capable of binding to the T cell CD3 receptor complex comprises a VL region selected from the group consisting of a VL region as depicted in SEQ ID NO: 35, 39, 125, 129, 161 or 165 of WO 2008/119567.

It is alternatively preferred that the second binding domain capable of binding to the T cell CD3 receptor complex comprises a VH region selected from the group consisting of a VH region as depicted in SEQ ID NO: 15, 19, 33, 37, 51, 55, 69, 73, 87, 91, 105, 109, 123, 127, 141, 145, 159, 163, 177 or 181 of WO 2008/119567.

More preferably, the CD33 targeting compound described herein is characterized by the second binding domain capable of binding to the T cell CD3 receptor complex comprising a VL region and a VH region selected from the group consisting of:

(a) a VL region as depicted in SEQ ID NO: 17 or 21 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 15 or 19 of WO 2008/119567;

(b) a VL region as depicted in SEQ ID NO: 35 or 39 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 33 or 37 of WO 2008/119567;

(c) a VL region as depicted in SEQ ID NO: 53 or 57 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 51 or 55 of WO 2008/119567;

5 (d) a VL region as depicted in SEQ ID NO: 71 or 75 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 69 or 73 of WO 2008/119567;

(e) a VL region as depicted in SEQ ID NO: 89 or 93 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 87 or 91 of WO 2008/119567;

10 (f) a VL region as depicted in SEQ ID NO: 107 or 111 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 105 or 109 of WO 2008/119567;

(g) a VL region as depicted in SEQ ID NO: 125 or 129 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 123 or 127 of WO 2008/119567;

(h) a VL region as depicted in SEQ ID NO: 143 or 147 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 141 or 145 of WO 2008/119567;

15 (i) a VL region as depicted in SEQ ID NO: 161 or 165 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 159 or 163 of WO 2008/119567; and

(j) a VL region as depicted in SEQ ID NO: 179 or 183 of WO 2008/119567 and a VH region as depicted in SEQ ID NO: 177 or 181 of WO 2008/119567.

20 According to a preferred embodiment of the CD33 targeting compound described herein, in particular the second binding domain capable of binding to the T cell CD3 receptor complex, the pairs of VH-regions and VL-regions are in the format of a single chain antibody (scFv). The VH and VL regions are arranged in the order VH-VL or VL-VH. It is preferred that the VH-region is positioned N-terminally to a linker sequence. The VL-region is positioned C-terminally of the linker sequence.

25 A preferred embodiment of the CD33 targeting compound described herein is characterized by the second binding domain capable of binding to the T cell CD3 receptor complex comprising an amino acid sequence selected from the group consisting of SEQ ID NOs: 23, 25, 41, 43, 59, 61, 77, 79, 95, 97, 113, 115, 131, 133, 149, 151, 167, 169, 185 or 187 of WO 2008/119567.

30 The formulations described herein are useful as pharmaceutical compositions in the treatment, amelioration and/or prevention of the pathological medical condition as described herein in a patient in need thereof. The term "treatment" refers to both therapeutic treatment and prophylactic or preventative measures. Treatment includes the application or administration of the formulation to the body, an isolated tissue, or cell from a patient who
35 has a disease/disorder, a symptom of a disease/disorder, or a predisposition toward a disease/disorder, with the purpose to cure, heal, alleviate, relieve, alter, remedy, ameliorate,

improve, or affect the disease, the symptom of the disease, or the predisposition toward the disease.

Those "in need of treatment" include those already with the disorder, as well as those in which the disorder is to be prevented. The term "disease" is any condition that would benefit from treatment with the protein formulation described herein. This includes chronic and acute disorders or diseases including those pathological conditions that predispose the mammal to the disease in question. Non-limiting examples of diseases/disorders to be treated herein include the herein described myeloid leukemia.

In some embodiments, the invention provides a pharmaceutical composition comprising a therapeutically effective amount of one or a plurality of the antibody construct described herein together with a pharmaceutically effective diluents, carrier, solubilizer, emulsifier, preservative, and/or adjuvant. Pharmaceutical compositions described herein include, but are not limited to, liquid, frozen, and lyophilized compositions.

Preferably, formulation materials are nontoxic to recipients at the dosages and concentrations employed.

In certain embodiments, the pharmaceutical composition may contain formulation materials for modifying, maintaining or preserving, for example, the pH, osmolarity, viscosity, clarity, color, isotonicity, odor, sterility, stability, rate of dissolution or release, adsorption or penetration of the composition. In such embodiments, suitable formulation materials include, but are not limited to, amino acids (such as glycine, glutamine, asparagine, arginine, proline, or lysine); antimicrobials; antioxidants (such as ascorbic acid, sodium sulfite or sodium hydrogen-sulfite); buffers (such as borate, bicarbonate, Tris-HCl, citrates, phosphates or other organic acids); bulking agents (such as mannitol or glycine); chelating agents (such as ethylenediamine tetraacetic acid (EDTA)); complexing agents (such as caffeine, polyvinylpyrrolidone, beta-cyclodextrin or hydroxypropyl-beta-cyclodextrin); fillers; monosaccharides; disaccharides; and other carbohydrates (such as glucose, mannose or dextrans); proteins (such as serum albumin, gelatin or immunoglobulins); coloring, flavoring and diluting agents; emulsifying agents; hydrophilic polymers (such as polyvinylpyrrolidone); low molecular weight polypeptides; salt-forming counterions (such as sodium); preservatives (such as benzalkonium chloride, benzoic acid, salicylic acid, thimerosal, phenethyl alcohol, methylparaben, propylparaben, chlorhexidine, sorbic acid or hydrogen peroxide); solvents (such as glycerin, propylene glycol or polyethylene glycol); sugar alcohols (such as mannitol or sorbitol); suspending agents; surfactants or wetting agents (such as pluronics, PEG, sorbitan esters, polysorbates such as polysorbate 20, polysorbate, triton, tromethamine, lecithin, cholesterol, tyloxapal); stability enhancing agents (such as sucrose or sorbitol);

tonicity enhancing agents (such as alkali metal halides, preferably sodium or potassium chloride, mannitol sorbitol); delivery vehicles; diluents; excipients and/or pharmaceutical adjuvants. See, REMINGTON'S PHARMACEUTICAL SCIENCES, 18th Edition, (A. R. Genmo, ed.), 1990, Mack Publishing Company.

- 5 In certain embodiments, the optimal pharmaceutical composition will be determined by one skilled in the art depending upon, for example, the intended route of administration, delivery format and desired dosage. See, for example, REMINGTON'S PHARMACEUTICAL SCIENCES, *supra*. In certain embodiments, such compositions may influence the physical state, stability, rate of in vivo release and rate of in vivo clearance of the antigen binding
- 10 proteins described herein. In certain embodiments, the primary vehicle or carrier in a pharmaceutical composition may be either aqueous or non-aqueous in nature. For example, a suitable vehicle or carrier may be water for injection, physiological saline solution or artificial cerebrospinal fluid, possibly supplemented with other materials common in compositions for parenteral administration. Neutral buffered saline or saline mixed with
- 15 serum albumin are further exemplary vehicles. In specific embodiments, pharmaceutical compositions comprise Tris buffer of about pH 7.0-8.5, or acetate buffer of about pH 4.0-5.5, and may further include sorbitol or a suitable substitute therefore. In certain embodiments of the invention, human antibody or antigen binding fragment thereof described herein or the antibody construct described herein compositions may be prepared for storage by mixing the
- 20 selected composition having the desired degree of purity with optional formulation agents (REMINGTON'S PHARMACEUTICAL SCIENCES, *supra*) in the form of a lyophilized cake or an aqueous solution. Further, in certain embodiments, the human antibody or antigen binding fragment thereof described herein or the antibody construct described herein may be formulated as a lyophilizate using appropriate excipients such as sucrose.
- 25 The pharmaceutical compositions described herein can be selected for parenteral delivery. Alternatively, the compositions may be selected for inhalation or for delivery through the digestive tract, such as orally. Preparation of such pharmaceutically acceptable compositions is within the skill of the art. The formulation components are present preferably in concentrations that are acceptable to the site of administration. In certain embodiments,
- 30 buffers are used to maintain the composition at physiological pH or at a slightly lower pH, typically within a pH range of from about 5 to about 8.

When parenteral administration is contemplated, the therapeutic compositions for use in this invention may be provided in the form of a pyrogen-free, parenterally acceptable aqueous solution comprising the desired human antibody or antigen binding fragment thereof

35 described herein or the antibody construct described herein in a pharmaceutically acceptable vehicle. A particularly suitable vehicle for parenteral injection is sterile distilled

water in which the antibody construct described herein is formulated as a sterile, isotonic solution, properly preserved. In certain embodiments, the preparation can involve the formulation of the desired molecule with an agent, such as injectable microspheres, bio-erodible particles, polymeric compounds (such as polylactic acid or polyglycolic acid), beads or liposomes, that may provide controlled or sustained release of the product which can be delivered via depot injection. In certain embodiments, hyaluronic acid may also be used, having the effect of promoting sustained duration in the circulation. In certain embodiments, implantable drug delivery devices may be used to introduce the desired antigen binding protein.

Additional pharmaceutical compositions will be evident to those skilled in the art, including formulations involving the antibody construct described herein in sustained- or controlled-delivery formulations. Techniques for formulating a variety of other sustained- or controlled-delivery means, such as liposome carriers, bio-erodible microparticles or porous beads and depot injections, are also known to those skilled in the art. See, for example, International Patent Application No. PCT/US93/00829, which is incorporated by reference and describes controlled release of porous polymeric microparticles for delivery of pharmaceutical compositions. Sustained-release preparations may include semipermeable polymer matrices in the form of shaped articles, e.g., films, or microcapsules. Sustained release matrices may include polyesters, hydrogels, polylactides (as disclosed in U.S. Pat. No. 3,773,919 and European Patent Application Publication No. EP 058481, each of which is incorporated by reference), copolymers of L-glutamic acid and gamma ethyl-L-glutamate (Sidman et al., 1983, Biopolymers 2:547-556), poly (2-hydroxyethyl-methacrylate) (Langer et al., 1981, J. Biomed. Mater. Res. 15:167-277 and Langer, 1982, Chem. Tech. 12:98-105), ethylene vinyl acetate (Langer et al., 1981, supra) or poly-D(-)-3-hydroxybutyric acid (European Patent Application Publication No. EP 133,988). Sustained release compositions may also include liposomes that can be prepared by any of several methods known in the art. See, e.g., Eppstein et al., 1985, Proc. Natl. Acad. Sci. U.S.A. 82:3688-3692; European Patent Application Publication Nos. EP 036,676; EP 088,046 and EP 143,949, incorporated by reference.

Pharmaceutical compositions used for in vivo administration are typically provided as sterile preparations. Sterilization can be accomplished by filtration through sterile filtration membranes. When the composition is lyophilized, sterilization using this method may be conducted either prior to or following lyophilization and reconstitution. Compositions for parenteral administration can be stored in lyophilized form or in a solution. Parenteral compositions generally are placed into a container having a sterile access port, for example,

an intravenous solution bag or vial having a stopper pierceable by a hypodermic injection needle.

Aspects of the invention includes self-buffering antibody construct formulations, which can be used as pharmaceutical compositions, as described in international patent application
5 WO 2006/138181A2 (PCT/US2006/022599), which is incorporated by reference in its entirety herein.

As discussed above, certain embodiments involve antibody construct protein compositions, particularly pharmaceutical compositions of the invention, that comprise, in addition to the antibody construct described herein, one or more excipients such as those illustratively
10 described in this section and elsewhere herein. Excipients can be used in the invention in this regard for a wide variety of purposes, such as adjusting physical, chemical, or biological properties of formulations, such as adjustment of viscosity, and or processes to improve effectiveness and or to stabilize such formulations and processes against degradation and spoilage due to, for instance, stresses that occur during manufacturing, shipping, storage,
15 pre-use preparation, administration, and thereafter.

A variety of expositions are available on protein stabilization and formulation materials and methods useful in this regard, such as Arakawa et al., "Solvent interactions in pharmaceutical formulations," Pharm Res. 8(3): 285-91 (1991); Kendrick et al., "Physical stabilization of proteins in aqueous solution," in: RATIONAL DESIGN OF STABLE PROTEIN
20 FORMULATIONS: THEORY AND PRACTICE, Carpenter and Manning, eds. Pharmaceutical Biotechnology. 13: 61-84 (2002), and Randolph et al., "Surfactant-protein interactions," Pharm Biotechnol. 13: 159-75 (2002), each of which is herein incorporated by reference in its entirety, particularly in parts pertinent to excipients and processes of the same for self-buffering protein formulations in accordance with the current invention,
25 especially as to protein pharmaceutical products and processes for veterinary and/or human medical uses.

Salts may be used in accordance with certain embodiments of the invention to, for example, adjust the ionic strength and/or the isotonicity of a formulation and/or to improve the solubility and/or physical stability of a protein or other ingredient of a composition in accordance with
30 the invention.

As is well known, ions can stabilize the native state of proteins by binding to charged residues on the protein's surface and by shielding charged and polar groups in the protein and reducing the strength of their electrostatic interactions, attractive, and repulsive interactions. Ions also can stabilize the denatured state of a protein by binding to, in
35 particular, the denatured peptide linkages ($--CONH$) of the protein. Furthermore, ionic

interaction with charged and polar groups in a protein also can reduce intermolecular electrostatic interactions and, thereby, prevent or reduce protein aggregation and insolubility.

Ionic species differ significantly in their effects on proteins. A number of categorical rankings of ions and their effects on proteins have been developed that can be used in formulating pharmaceutical compositions in accordance with the invention. One example is the Hofmeister series, which ranks ionic and polar non-ionic solutes by their effect on the conformational stability of proteins in solution. Stabilizing solutes are referred to as "kosmotropic." Destabilizing solutes are referred to as "chaotropic." Kosmotropes commonly are used at high concentrations (e.g., >1 molar ammonium sulfate) to precipitate proteins from solution ("salting-out"). Chaotropes commonly are used to denature and/or to solubilize proteins ("salting-in"). The relative effectiveness of ions to "salt-in" and "salt-out" defines their position in the Hofmeister series.

Free amino acids can be used in the antibody construct formulations in accordance with various embodiments of the invention as bulking agents, stabilizers, and antioxidants, as well as other standard uses. Lysine, proline, serine, and alanine can be used for stabilizing proteins in a formulation. Glycine is useful in lyophilization to ensure correct cake structure and properties. Arginine may be useful to inhibit protein aggregation, in both liquid and lyophilized formulations. Methionine is useful as an antioxidant.

Polyols include sugars, e.g., mannitol, sucrose, and sorbitol and polyhydric alcohols such as, for instance, glycerol and propylene glycol, and, for purposes of discussion herein, polyethylene glycol (PEG) and related substances. Polyols are kosmotropic. They are useful stabilizing agents in both liquid and lyophilized formulations to protect proteins from physical and chemical degradation processes. Polyols also are useful for adjusting the tonicity of formulations.

Among polyols useful in select embodiments of the invention is mannitol, commonly used to ensure structural stability of the cake in lyophilized formulations. It ensures structural stability to the cake. It is generally used with a lyoprotectant, e.g., sucrose. Sorbitol and sucrose are among preferred agents for adjusting tonicity and as stabilizers to protect against freeze-thaw stresses during transport or the preparation of bulks during the manufacturing process.

Reducing sugars (which contain free aldehyde or ketone groups), such as glucose and lactose, can glycate surface lysine and arginine residues. Therefore, they generally are not among preferred polyols for use in accordance with the invention. In addition, sugars that form such reactive species, such as sucrose, which is hydrolyzed to fructose and glucose under acidic conditions, and consequently engenders glycation, also is not among preferred

polyols of the invention in this regard. PEG is useful to stabilize proteins and as a cryoprotectant and can be used in the invention in this regard.

Embodiments of the antibody construct formulations further comprise surfactants. Protein molecules may be susceptible to adsorption on surfaces and to denaturation and consequent aggregation at air-liquid, solid-liquid, and liquid-liquid interfaces. These effects generally scale inversely with protein concentration. These deleterious interactions generally scale inversely with protein concentration and typically are exacerbated by physical agitation, such as that generated during the shipping and handling of a product.

Surfactants routinely are used to prevent, minimize, or reduce surface adsorption. Useful surfactants in the invention in this regard include polysorbate 20, polysorbate 80, other fatty acid esters of sorbitan polyethoxylates, and poloxamer 188.

Surfactants also are commonly used to control protein conformational stability. The use of surfactants in this regard is protein-specific since, any given surfactant typically will stabilize some proteins and destabilize others.

Polysorbates are susceptible to oxidative degradation and often, as supplied, contain sufficient quantities of peroxides to cause oxidation of protein residue side-chains, especially methionine. Consequently, polysorbates should be used carefully, and when used, should be employed at their lowest effective concentration. In this regard, polysorbates exemplify the general rule that excipients should be used in their lowest effective concentrations.

Embodiments of the antibody construct formulations further comprise one or more antioxidants. To some extent deleterious oxidation of proteins can be prevented in pharmaceutical formulations by maintaining proper levels of ambient oxygen and temperature and by avoiding exposure to light. Antioxidant excipients can be used as well to prevent oxidative degradation of proteins. Among useful antioxidants in this regard are reducing agents, oxygen/free-radical scavengers, and chelating agents. Antioxidants for use in therapeutic protein formulations in accordance with the invention preferably are water-soluble and maintain their activity throughout the shelf life of a product. EDTA is a preferred antioxidant in accordance with the invention in this regard.

Antioxidants can damage proteins. For instance, reducing agents, such as glutathione in particular, can disrupt intramolecular disulfide linkages. Thus, antioxidants for use in the invention are selected to, among other things, eliminate or sufficiently reduce the possibility of themselves damaging proteins in the formulation.

Formulations in accordance with the invention may include metal ions that are protein co-factors and that are necessary to form protein coordination complexes, such as zinc

necessary to form certain insulin suspensions. Metal ions also can inhibit some processes that degrade proteins. However, metal ions also catalyze physical and chemical processes that degrade proteins.

Magnesium ions (10-120 mM) can be used to inhibit isomerization of aspartic acid to isoaspartic acid. Ca^{+2} ions (up to 100 mM) can increase the stability of human deoxyribonuclease. Mg^{+2} , Mn^{+2} , and Zn^{+2} , however, can destabilize rhDNase. Similarly, Ca^{+2} and Sr^{+2} can stabilize Factor VIII, it can be destabilized by Mg^{+2} , Mn^{+2} and Zn^{+2} , Cu^{+2} and Fe^{+2} , and its aggregation can be increased by Al^{+3} ions.

Embodiments of the antibody construct formulations further comprise one or more preservatives. Preservatives are necessary when developing multi-dose parenteral formulations that involve more than one extraction from the same container. Their primary function is to inhibit microbial growth and ensure product sterility throughout the shelf-life or term of use of the drug product. Commonly used preservatives include benzyl alcohol, phenol and m-cresol. Although preservatives have a long history of use with small-molecule parenterals, the development of protein formulations that includes preservatives can be challenging. Preservatives almost always have a destabilizing effect (aggregation) on proteins, and this has become a major factor in limiting their use in multi-dose protein formulations. To date, most protein drugs have been formulated for single-use only. However, when multi-dose formulations are possible, they have the added advantage of enabling patient convenience, and increased marketability. A good example is that of human growth hormone (hGH) where the development of preserved formulations has led to commercialization of more convenient, multi-use injection pen presentations. At least four such pen devices containing preserved formulations of hGH are currently available on the market. Norditropin (liquid, Novo Nordisk), Nutropin AQ (liquid, Genentech) & Genotropin (lyophilized--dual chamber cartridge, Pharmacia & Upjohn) contain phenol while Somatropin (Eli Lilly) is formulated with m-cresol. Several aspects need to be considered during the formulation and development of preserved dosage forms. The effective preservative concentration in the drug product must be optimized. This requires testing a given preservative in the dosage form with concentration ranges that confer anti-microbial effectiveness without compromising protein stability.

As might be expected, development of liquid formulations containing preservatives are more challenging than lyophilized formulations. Freeze-dried products can be lyophilized without the preservative and reconstituted with a preservative containing diluent at the time of use. This shortens the time for which a preservative is in contact with the protein, significantly minimizing the associated stability risks. With liquid formulations, preservative effectiveness and stability should be maintained over the entire product shelf-life (about 18 to 24 months).

An important point to note is that preservative effectiveness should be demonstrated in the final formulation containing the active drug and all excipient components.

The antibody construct described herein generally will be designed for specific routes and methods of administration, for specific administration dosages and frequencies of administration, for specific treatments of specific diseases, with ranges of bio-availability and persistence, among other things. Formulations thus may be designed in accordance with the invention for delivery by any suitable route, including but not limited to orally, aurally, ophthalmically, rectally, and vaginally, and by parenteral routes, including intravenous and intraarterial injection, intramuscular injection, and subcutaneous injection.

Once the pharmaceutical composition has been formulated, it may be stored in sterile vials as a solution, suspension, gel, emulsion, solid, crystal, or as a dehydrated or lyophilized powder. Such formulations may be stored either in a ready-to-use form or in a form (e.g., lyophilized) that is reconstituted prior to administration. The invention also provides kits for producing a single-dose administration unit. The kits of the invention may each contain both a first container having a dried protein and a second container having an aqueous formulation. In certain embodiments of this invention, kits containing single and multi-chambered pre-filled syringes (e.g., liquid syringes and lyosyringes) are provided. The therapeutically effective amount of an antibody construct protein-containing pharmaceutical composition to be employed will depend, for example, upon the therapeutic context and objectives. One skilled in the art will appreciate that the appropriate dosage levels for treatment will vary depending, in part, upon the molecule delivered, the indication for which the antibody construct described herein is being used, the route of administration, and the size (body weight, body surface or organ size) and/or condition (the age and general health) of the patient. In certain embodiments, the clinician may titer the dosage and modify the route of administration to obtain the optimal therapeutic effect. A typical dosage may range from about 0.1 µg/kg to up to about 30 mg/kg or more, depending on the factors mentioned above. In specific embodiments, the dosage may range from 1.0 µg/kg up to about 20 mg/kg, optionally from 10 µg/kg up to about 10 mg/kg or from 100 µg/kg up to about 5 mg/kg.

A therapeutically effective amount of pharmaceutical composition of the invention comprising the CD33 targeting compound and at least one epigenetic factor in a single or separate formulations preferably results in a decrease in severity of disease symptoms, in increase in frequency or duration of disease symptom-free periods or a prevention of impairment or disability due to the disease affliction. For treating CD33-expressing tumors, a therapeutically effective amount of the CD33 targeting compound and at least one epigenetic factor in a single or separate formulations, e.g. an anti-CD33/CD3 antibody construct and the

at least one epigenetic factor, preferably inhibits cell growth or tumor growth by at least about 20%, at least about 40%, at least about 50%, at least about 60%, at least about 70%, at least about 80%, or at least about 90% relative to untreated patients. The ability of a compounds to inhibit tumor growth may be evaluated in an animal model predictive of efficacy in human tumors.

Pharmaceutical compositions may be administered using a medical device. Examples of medical devices for administering pharmaceutical compositions are described in U.S. Patent Nos. 4,475,196; 4,439,196; 4,447,224; 4,447, 233; 4,486,194; 4,487,603; 4,596,556; 4,790,824; 4,941,880; 5,064,413; 5,312,335; 5,312,335; 5,383,851; and 5,399,163, all incorporated by reference herein.

It should be understood that the inventions herein are not limited to particular methodology, protocols, or reagents, as such can vary. The discussion and examples provided herein are presented for the purpose of describing particular embodiments only and are not intended to limit the scope of the present invention, which is defined solely by the claims.

All publications and patents cited throughout the text of this specification (including all patents, patent applications, scientific publications, manufacturer's specifications, instructions, etc.), whether supra or infra, are hereby incorporated by reference in their entirety. Nothing herein is to be construed as an admission that the invention is not entitled to antedate such disclosure by virtue of prior invention. To the extent the material incorporated by reference contradicts or is inconsistent with this specification, the specification will supersede any such material.

Examples:

The following examples are provided for the purpose of illustrating specific embodiments or features of the present invention. These examples should not be construed as to limit the scope of this invention. The examples are included for purposes of illustration, and the present invention is limited only by the claims.

Example 1 – Epigenetic modifying drugs as sensitizing agents for CD33 targeting compound-induced cytotoxicity

The level of CD33 expression was identified as a critical variable for the extent of activity of a CD33 targeting compound described herein such as AMG330 against human AML cells. Accordingly, it was surprising to observe the potential of epigenetic modifying drugs such as histone deacetylase (HDAC) inhibitors or DNA methyltransferase (DNMT) I inhibitors as sensitizing agents for AMG 330-induced cytotoxicity.

For the experimental settings mononuclear cells were collected from healthy adult volunteers via leukapheresis, and T cells enriched through magnetic cell sorting (Pan T Cell Isolation Kit II; Miltenyi Biotec, Auburn, CA). Thawed cell aliquots were labeled with 3 μ M CellVue Burgundy (eBioscience, San Diego, CA) according to the manufacturer's instructions.

- 5 Human myeloid OCI-AML3, KG-1a, ML-1, NB4, TF-1, and HL-60 cells were maintained as previously described (Walter et al. Blood. 2003;102(4):1466-1473; Walter et al. Blood. 2004;103(11):4276-4284, Walter et al. Blood. 2005;105(3):1295-1302).

CD33 expression on parental AML cells and cell lines was quantified by flow cytometry using a phycoerythrin (PE)-conjugated anti-CD33 antibody (clone P67.6; BD Biosciences, San
10 Jose, CA, USA) (see Walter et al. Blood. 2005;105(3):1295-1302).

For the quantification of compound induces cytotoxicity AML cells were taken during exponential growth and incubated at 37°C (in 5% CO₂ and air) in 96-well round bottom plates (BD Falcon™) at 5-10 x 10³ cells/well in 225 μ L culture medium containing various concentrations of AMG 330 as well as T-cells at different effector:target (E:T) cell ratios.
15 After 48 hours, cell numbers and drug-induced cytotoxicity, using 4',6-diamidino-2-phenylindole (DAPI) to detect non-viable cells, were determined using a LSRII flow cytometer (BD Biosciences, San Jose, CA) and analyzed with FlowJo (Tree Star, Ashland, OR). AML cells were identified by forward/side scatter properties and negativity for CellVue Burgundy dye.

20 To determine CD33 modulation, aliquots of AML cells were left untreated or incubated with either AMG 330 (250 pg/mL) or an unconjugated, unlabeled anti-CD33 antibody (clone P67.6; BD Biosciences, San Jose, CA, USA; 2.5 μ g/mL). After 48 hours, cells were washed in ice-cold Phosphate Buffered Saline (PBS, GIBCO Invitrogen) to remove unbound antibody and resuspended in PBS containing 2% fetal bovine serum. As AMG 330 does not compete
25 for binding to CD33 with P67.6, aliquots of untreated and AMG 330-treated cells were incubated with P67.6 or no primary antibody followed by biotin-conjugated rat anti-mouse IgG₁ (used at 2.5 μ g/mL in PBS/2% FBS) and streptavidin-phycoerythrin (PE) (used at 2.5 μ g/mL in PBS/2% FBS: both from BD Biosciences). Likewise, aliquots of untreated and P67.6-treated cells were incubated with unconjugated P67.6 or without primary antibody;
30 cells were then washed and incubated with biotin-conjugated rat anti-mouse IgG₁ monoclonal antibody followed by incubation with streptavidin-PE conjugate. To identify nonviable cells, all samples were stained with DAPI. At least 10,000 events were acquired, and DAPI⁺ cells were analyzed on a Canto flow cytometer (BD Biosciences) using FlowJo Software. Linear median fluorescence intensity (MFI) values were used to calculate the
35 percentage of drug-bound CD33 internalization.

As shown in Figure 1, 3-day pretreatment with the HDAC inhibitor, panobinostat, resulted in significant increase in CD33 expression in OCI-AML3 and, markedly, KG-1a cells. More importantly, relative to untreated cells, cells pretreated with panobinostat for 3 days were modestly more sensitive to AMG 330-induced cytotoxicity (KG-1a > OCI-AML3).

Pretreatment with the DNMT I inhibitor, azacitidine, for 3 days resulted in significant increase in CD33 expression on KG-1a. Consistently, after pretreatment with azacitidine, KG-1a cells became significantly more sensitive to AMG 330-induced cytotoxicity relative to untreated cells (Figure 2).

Example 2 – CD33 up-regulation on AML cells to increase AMG 330 mediated lysis efficacy

Hydroxyurea

Up-regulation of CD33 on AML cell lines:

AML cell lines HL-60, PL21, OCI-AML3, KG1a and MV4-11 were seeded in 24-well plates at 1×10^6 cells/ml on day 0. Cells were either left untreated (UT) or treated with 10 μ M (H1) or 100 μ M (H2) hydroxyurea (Sigma) for three consecutive days (day 0, day 1 and day 2). On day 3, cells were harvested, counted and analyzed for changes in CD33 surface expression level by flow cytometry.

Figure 3 and Table 2 show an upregulation of CD33 on HL-60 and PL21 AML cells in a concentration dependant manner.

CD33 MFI Ratio	UT	H1	H2
HL-60	134.9	171.3	210.0
PL21	166.9	177.9	191.8

Table 2: CD33 MFI Ratios of HL-60 and PL21 AML cells after incubation with/without hydroxyurea determined by flow cytometry.

Up-regulation of CD33 on primary AML cells:

Three primary AML samples were seeded in a 12-well plates at 1×10^6 cells/ml, as previously described (Krupka, Subklewe et.al, Blood 2014). Cells were either left untreated (UT) or treated with 10 μ M (H1) or 100 μ M (H2) hydroxyurea (Sigma) for three consecutive days (day 0, day 1 and day 2). On day 1 - day 3, cells were harvested, counted and analyzed for changes in CD33 surface expression by flow cytometry.

Figure 4 and Table 3 show an up-regulation of CD33 in 1 of 3 patient samples on day 1 and day 2 of incubation with hydroxyurea.

CD33 MFI Ratio	UT	H1	H2
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day 1 Patient 2	3.7	4.2	4.5
day 2 Patient 2	4.8	4.2	5.1

Table 3: CD33 MFI Ratios of primary AML cells from patient 2 after 1 and 2 days of incubation with hydroxyurea determined by flow cytometry.

5 Granulocyte-Colony Stimulating Factor (G-CSF)

Up-regulation of CD33 on AML cell lines:

AML cell lines OCI-AML3 and KG1 α were seeded in 24-well plates at 5×10^5 cells/ml on day 0. Cells were either left untreated (UT) or treated with 200 ng/ml (C1) or 2000 ng/ml (C2) G-CSF (Peprotech) for 10 days. G-CSF was freshly added to the cultures biweekly. On day 10, cells were harvested and analyzed for changes in CD33 surface expression by flow cytometry.

Figure 5 and Table 4 show a low upregulation of CD33 on KG1 α AML cells in an concentration dependant manner.

CD33 MFI Ratio	UT	C1	C2
KG1 α	16.5	17.9	19.2

- 15 Table 4: CD33 MFI Ratios of KG1 α AML cells after 10 days of incubation with/without G-CSF determined by flow cytometry.

SEQ ID NO.	DESIGNATION	SOURCE	TYPE	SEQUENCE
1	CD33 VH of AH3	artificial	aa	QVQLVQSGAEVKKPGESVKVSKKASGYTFITNYGMNWRQAPGQGLEWMGWINTYTGEPYADDFKG RVTMSDDTSTSTAYLEINSLRSDDTAIYYCARWSWSDGYVYFDYWGQGTTVTVSS
2	CD33 HCDR1 of AH3	artificial	aa	NYGMN
3	CD33 HCDR2 of AH3	artificial	aa	WINTYTGEPYADDFKG
4	CD33 HCDR3 of AH3	artificial	aa	WSWSDGYVYFDY
5	CD33 VH of AH3	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAGGTAAGGCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAGGAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAACACCTACACTGGAGAGCCAAATATGCTGATGACTTCAAGGGA CGGGTTACCATGCTTCGGATACCTCTACCAGCACTGCCATTTGGAAATCAACAGCCTCAGAAAT GATGACACGGCTATATATATCTGTGCGCGCTGGAGTTGGAGTGTGCTTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCACCTACGGTCACCGTCTCCTCA
6	CD33 VL of AH3	artificial	aa	DIVMTQSPDLSITVSLGERITINCKSSQSVLDSKKNKNSLAWYQQKPGQPPKILLISWASTRESGIPD RFGSGSGTDFTLTIDSLQPEDSATYFCQQSAHFPIITFGQGRLEIK
7	CD33 LCDR1 of AH3	artificial	aa	KSSQSVLDSKKNKNSLA
8	CD33 LCDR2 of AH3	artificial	aa	WASTRES
9	CD33 LCDR3 of AH3	artificial	aa	QQSAHFPIIT
10	CD33 VL of AH3	artificial	nt	GACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGGCGAGAGGACCAACATCAAC TGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAGAAATAAGAACTCCTTAGCTTGGTACCAGCAG AAACCCAGGACAGCCTCCCTAAATTAATCTCTTCTTGGGATCTACGCGGAAATCCGGGATCCCTGAC CGATTCAAGTGGCAGCGGGTCTGGGACAGATTTCATCTCATTTATGACAGCCTGCAGCCTGAAGAT TCTGCAACTTACTATTTGTCAACAGTCTGCCCACTTCCCGATCACCTTTGGCCAAAGGACACGACTG GAGATTAAA
11	CD33 CD33 HL of AH3	artificial	aa	QVQLVQSGAEVKKPGESVKVSKKASGYTFITNYGMNWRQAPGQGLEWMGWINTYTGEPYADDFKG RVTMSDDTSTSTAYLEINSLRSDDTAIYYCARWSWSDGYVYFDYWGQGTTVTVSSGGGSGGGGS GGGSDIVMTQSPDLSITVSLGERITINCKSSQSVLDSKKNKNSLAWYQQKPGQPPKILLISWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYFCQQSAHFPIITFGQGRLEIK
12	CD33 HL of AH3	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAGGTAAGGCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAGGAGGCTCCAGGACAGGGT

13	CD33 AH3 HL x H2C HL	artificial	aa	TTAGAGTGGATGGGCTGGATAAAACACCTACACTGGAGAGCCAAACATATGCTGATGACTTCAAGGGA CGGGTTACCATGTCTTCGGATACCTCTACCAGCACTGCCTATTTGGAAATCAACAGCCTCAGAAAT GATGACACGGCTATATATTAATGCTGCGCGCTGGAGTTGGAGTGATGTTACTACGTTTACTTTTGAC TACTGGGGCCAAAGCACTACGGTCACCGTCTCCTCAGGTGGTGGTGTCTTGGCGGCGGCTCC GGTGGTGGTGGTCTGACATCGTGATGACACAGTCTCCAGACTCCTCAGCTGTGTCTCTGGCGGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTCTTAAATTAATCTCCTTCTGGCATCTACGCGGAA GCTTGGTACAGAGAAACAGGACAGCTCCTAAATTAATCTCCTTCTGGCATCTACGCGGAA TCCGGGATCCCTGACCGGATTCAGTGGCAGCGGTCTGGGACAGATTCACCTTCCCGATCACCTTTGGC CTGACGCTGAAAGATTCGCAACTTACTATTTGTTCAACAGTCTGCCCATCTTCCCGATCACCTTTGGC CAAGGACACGACTGGAGATTAAA QVQLVQSGAEVKKPESVKVSKASGYFTNYGMNWRQAPGQGLEWMGWINTYTGEPYADDFKG RVTMSDDTSTSTAYLEINSLRSDDTAIYYCARWSWSDGYVYFDYWGQGTITVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSVLDSKKNKNSLAWYQQKPGQPCKLLLSWASTRE SGIPDRFSGSGGTDFLTITIDSLQPEDSATYYCQQSAHFPITFGQGTREIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNWRQAPGKLEWVARIRSKYNNYATYYADSVKDRFTISR DDSKNTAYLQMNLIKTEDTAVYCVRHGNFGNSYISYWAYWGQGTITVTVSSGGGSGGGGSGGGGS QTVVTQEPSTLVSPGGTITLTGSSSTGAVTSGYYPNWQQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEFYCALMYSNRWVFGGKLTIVL
14	CD33 AH3 HL x H2C HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAAAAGCCTGGAGAGTCAGTCAAGGTCTCTCTGC AAGGTAGCGGTGATACCTTCACAACTATGGAATGAACTGGGTGAGGAGGCTCCAGGACAGGT TTAGAGTGGATGGGCTGGATAAAACACCTACACTGGAGAGCCAAACATATGCTGATGACTTCAAGGGA CGGGTTACCATGTCTTCGGATACCTCTACCAGCACTGCCTATTTGGAAATCAACAGCCTCAGAAAT GATGACACGGCTATATATTAATGCTGCGCGCTGGAGTTGGAGTGATGTTACTACGTTTACTTTTGAC TACTGGGGCCAAAGCACTACGGTCACCGTCTCCTCAGGTGGTGGTGTCTTGGCGGCGGCTCC GGTGGTGGTGGTCTGACATCGTGATGACACAGTCTCCAGACTCCTCAGCTGTGTCTCTGGCGGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTCTTAAATTAATCTCCTTCTGGCATCTACGCGGAA GCTTGGTACAGAGAAACAGGACAGCTCCTAAATTAATCTCCTTCTGGCATCTACGCGGAA TCCGGGATCCCTGACCGGATTCAGTGGCAGCGGTCTGGGACAGATTCACCTTCCCGATCACCTTTGGC CTGACGCTGAAAGATTCGCAACTTACTATTTGTTCAACAGTCTGCCCATCTTCCCGATCACCTTTGGC CAAGGACACGACTGGAGATTAAAATCCGAGGTGGTGGCTCCGAGGTGAGTGGTGGATTCACCTTCAAT GAGTACGCCATGAACTGGGTCCGCCAGGTCCAGGAAAGGTTTGGAAATGGGTGTCTCGCATAAAGA AGTAAATATAATAATATGCAACATATTAATGCGCATTCAGTGAAGACAGGTTACCATCTCCAGA GATGATTCAAAAAACACTGCCATCTACAAATGAACAACTTGAAATGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACCTTCGGTAAATAGCTACATATCTTACTGGGTACTTGGGGCCAAAGG ACTTGGTCAACCGTCTCCTCAGGTGGTGGTGTCTTGGCGGCGGCGGTCCGGTGGTGGTGGTCT CAGACTGTGTGACTCAGGAACCTTCACCTACCGTATCACCTGGTGAACAGTCACTACCTTGT GGCTCCTCGACTGGGCTGTATACATCTGGCTACTACCCAACTGGGTCCACAAAAACAGGTCTAG GCACCCCGTGGTCTAATAGGTGGGACTAAGTTCTCCTCGCCCCGGTACTCCTGCCAGATTCTCAGGC

15	CD33 AH3 HL x F12Q HL	artificial	aa	TCCCTGCTTGGAGGCAAGGCTGCCCTCACCCCTCACAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGCTCTATGGTACAGCAACCGCTGGGTGTTCCGGTGGAGGAACCAAACTGACTGTCTCTA QVQLVQSGAEVVKPGEISVKVSKKASGYTFITNYGMNWVRQAPGQGLEWMGWINTYITGEPTYADDFKG RVTMSDDTSTAYLIEINLSRSDDTAIYYCARWSWSDGYVYFDYWQGGTITVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSLDSSKNKNSLAWYQQKPKPPKLLISWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYCYQQSAHFPITFTGQGRLEIKSGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSVKGRFTISR DDSKNTAYLQMNILKTEDTAVYYCVRHGNTGNSYVSWAYWQGTILTVSSGGGSGGGSGGGGS QTVVTPQEPSLTVSPGGTFTLTTCGSSTGAVTSGNYPNWVQOKPQAPRGLIGGTRKFLAPGTPARFSG SLLGGKAALTLISGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL
16	CD33 AH3 HL x F12Q HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAAAAGCCTGGAGAGTCAGTCAAGGTCTCTCTGC AAGCTAGCGGGTATACCTTCACAAAATATGGAAATGAACTGGGTGAGGCAGGCTCCAGGACAGGT TTAGAGTGGATGGGTGGATAAACACCTACACTGGAGAGCCAAATATGCTGATGACTTCAAGGGA CGGTTACCATGCTTTCGGATACCTCTACCCAGCACTGCCATATTTGAAAATCAACAGCCTCAGAAAT GATGACACGGCTATATATTTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGCGCAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTGTCTGGCGCGCGGCTCC GGTGGTGGTGTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAAGATAAGAACTCCCTTA GCTTGGTACCAGAGAAAACAGGACAGCTCTCTAAATTAATCTCTTCTGGGCATCTACGCGGAA TCCGGGATCCCTTGACCGATTCAGTGGCAGCGGTCTGGGACAGATTCACCTCTCACTATTTGACAGC CTGCAGCCTGAAAGATTCGCAACTTACTATTTGTCAACAGTCTGCCCCTTCCCGATCACCTTTGGC CAAGGACACAGCTGGAGATTAAATCCGGAGGTGGTGGCTCCGAGGTGACGTGGTTCGAGTCTGGA GGAGGATTGGTGCAGCTGGAGGGTCAATTGAAATCTCATGTGCAGCTCTGGATTCACCTTCAAT AGCTACGCCATGAATGGGTCCCGCAGGCTCCAGGAAAGGTTTGGAAATGGGTGCTCGCATAGA AGTAAATATAATTAATGCAACATATTTATGCCGATTCAGTGAAAGGCAGGTTACCATCTCCAGA GATGATTCAAAAACACTGCCATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAATTCGGTAATAGTACGTTTCTTGGTGGCTTACTGGGCGCAAGGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGTCTTGGCGCGCGGCTCCCGTGGTGGTGTCT CAGACTGTGTGACTCAGGAACCTTCACTACCGTATCACTGGTGGAACAGTCACACTCACTGT GGCTCCTCGACTGGGGCTGTACATCTGGCAACTACCCAACTGGGTCCAAACAAAAACAGGTCTCAG GCACCCCGTGGTCTTAATAGGTGGGACTAAAGTTCCCTCGCCCCCGGTACTCTTCCAGATTCCTCAGGC TCCCTGTCTGGAGGCAAGGTGCCCTCACCTCTCAGGGTGACAGGCAGGATGAGGACAGAAATAT TACTGTGTCTCTATGGTACAGCAACCGCTGGGTGTCTCGGTGGAGGAACCAAACTGACTGTCTCTA
17	CD33 AH3 HL x I2C HL	artificial	aa	QVQLVQSGAEVVKPGEISVKVSKKASGYTFITNYGMNWVRQAPGQGLEWMGWINTYITGEPTYADDFKG RVTMSDDTSTAYLIEINLSRSDDTAIYYCARWSWSDGYVYFDYWQGGTITVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSLDSSKNKNSLAWYQQKPKPPKLLISWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYCYQQSAHFPITFTGQGRLEIKSGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSVKDRFTISR DDSKNTAYLQMNILKTEDTAVYYCVRHGNTGNSYISWAYWQGTILTVSSGGGSGGGSGGGGS

18	CD33 AH3 HL x I2C HL	artificial	nt	QTVVTFEPSTLVSPGGTVTLTTCGSSTGAVTSGNYPNWVQQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGKKAALTLISGVQPEDEAEFYCVLWYSNPWFVGGGTKLTVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAGGACAGGCTCCAGGACAGGGT TTAGAGTGGATGGGCTGGATAAAACACCTACACTGGAGAGCAACATATGCTGATGACTTCAAGGGA CGGTTACCATGTCTTCGGATACCTCTACCAGCACTGCCTATTTGGAAATCAACAGCCTCAGAAAGT GATGACACGGCTATATATCTGTGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCAAGGCACCTACGGTCACCGCTCCTCAGGTGGTGGTTCTTGGCGGCGGCTCC GGTGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGCTCTGGGCGAG AGGACACCATCAACTGCAAGTCCAGCAGAGTGTCTTAGACAGCTCCAAAGATAAGAACTCCTTA GCTTGGTACACAGAGAAAACAGGACAGCTCCTAAAATTACTCTCTTCTTGGGCACTTACGCGGGA TCCGGGATCCCCTGACCGATTCTAGTGGCAGCGGTCTGGGACAGATTCTACTCTCACTATTGACAGC CTGCAGCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCAGATCACCTTTGGC CAAGGACACGACTGGAGATTAAATCCGAGGTGGTGGCTCCGAGGTGCAGCTGGTCGAGTCTGGA GGAGGATTGGTCAGCTGGAGGGTCAATTGAAACTCTCATGTGCAGCTCTGGATTCACTTCAAT AAGTACGCCATGAACTGGGTCCGCGAGGCTCCAGGAAAAGGTTTGGAAATGGGTGCTCGCATAGA AGTAAATATAAATAATTATGCAACATATTATGCCGATTTCAGTGAAGACAGGTTACCATCTCCAGA GATGATTCAAAAAACACTGCCTATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAATAGTACATATCTTACTGGGTACTTGGGGCAAGGG ACTGTGGTCAACCGTCTCCTCAGGTGGTGGTGTCTGGCGGCGGGCTCCGGTGGTGGTGTCT CAGACTGTTGTGACTCAGGAACTTCACTACCGTATCACTGGTGGAAACAGTCACTCACTTGT GGTCTCTCGACTGGGCTGTACATCTGGCAACTACCCAACTGGGTCCAAACAAACAGGTCAG GCACCCGCTGGTCTAAATAGGTGGACTAAGTCTCTCGCCCCGTACTCTTCCAGATTCTCAGGC TCCCTGCTTGGAGGCAAGGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGTCTTCTATGGTACAGCAACCGCTGGGTGTCTCGGTGGAGAAACCAACTGACTGTCTTA QVQLVQSGAEVKKPGASVKVSCKASGYTFITNYGMNWVKQAPGQGLKWMGWINTYTGEPTIYADDFKG RVTMTSDTSTSTAYLELHNLRSDDTAVYYCARWSWSVSDGYVYFDYWGQGTTITVTVSS NYGMN
19	CD33 VH of AF5	artificial	aa	QVQLVQSGAEVKKPGASVKVSCKASGYTFITNYGMNWVKQAPGQGLKWMGWINTYTGEPTIYADDFKG RVTMTSDTSTSTAYLELHNLRSDDTAVYYCARWSWSVSDGYVYFDYWGQGTTITVTVSS NYGMN
20	CD33 HCDR1 of AF5	artificial	aa	NYGMN
21	CD33 HCDR2 of AF5	artificial	aa	WINTYTGEPTIYADDFKG
22	CD33 HCDR3 of AF5	artificial	aa	WSWSVSDGYVYFDY
23	CD33 VH of AF5	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAAGCCTGGAGCGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAAACAGGCTCCAGGACAGGGT TTAAAGTGGATGGGCTGGATAAAACACCTACACTGGAGAGCAACATATGCTGATGACTTCAAGGGA CGGTTACCATGACTTCGGATACCTCTACCAGCACTGCCTATTTGGAACTCCACAACTCAGAAAGT

				GATGACACGGCTGTATATTACTGTGTGGCGCTGGAGTTGGAGTGTGGTTACTACGTTTACTTTTAC
24	CD33 VL of AF5	artificial	aa	TACTGGGGCCAAAGGCACACACGCTACCGGTACCGTCTCCTCA DIVMTQSPDRLTVSLGERTTINCKSSQSVLDSSKNKNSLAWYQQKPGQPPKLLLSWASTRESGIPD RFSGSGSGTDFTLTIDSLQPEDSATYICQQSAHFPITFGQGRLEIK KSSQSVLDSSKNKNSLA
25	CD33 LCDR1 of AF5	artificial	aa	
26	CD33 LCDR2 of AF5	artificial	aa	WASTRES
27	CD33 LCDR3 of AF5	artificial	aa	QQSAHFPIT
28	CD33VL of AF5	artificial	nt	GACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGGCGAGAGGACCACCATCAAC TGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCAAGAAATAAGAACTCCTTAGCTTGGTACACAGAG AAACACAGGACAGCTCCTAAATTTACTCCTTTCTTGGGCACTACGGGGAAATCCGGGATCCTCTGAC CGATTACGTGGCAGCGGTCTGGGACAGATTTTCACTCTCACTATTGACAGCTGCAGCTGAAAGAT TCTGTCAACTTACTATTGTCAACAGTCTGCCCACTTCCCGATCACCTTTGGCCAAAGGACACGACTG GAGATTAAA
29	CD33HL of AF5	artificial	aa	QVQLVQSGAEVKKPGASVKVSCKASGYTFITNYGMNWVKQAPGQGLKWMGWINTYTGEPYADDFKG RVMTSDTSTSTAYLELHNLRSDDTAVYICARWSWSDGYVYFDYWGQGTITVTVSSGGGGSGGGGS GGGSDIVMTQSPDRLTVSLGERTTINCKSSQSVLDSSKNKNSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYICQQSAHFPITFGQGRLEIK
30	CD33 HL of AF5	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCTTGAGCGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAAACTATGGAATGAATGAGTGAAGCAGGCTCCAGGACAGGCT TTAAAAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCAACATATGCTGATGACTTCAAGGGA CGGTTACCATGACTTCGGATACCTCTACACAGACTGCCTATTTGGAACCTCCACAACTTCAGAAAT GATGACACGGCTGTATATTACTGTGTGGCGCTGGAGTTGGAGTGTGATGTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCACACTACGGTCACCGTCTCCTCAGGTGGTGGTGTCTTGGCGGGCGGCTCC GGTGGTGGTGGTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTTGGCGGAG AGGACCACTCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCAAGAAATAAGAACTCCTTTA GCTTGGTACCAGAGAAACCAGGACAGCTCCTAAATTTACTCCTTTCTTGGGCACTACCGGGAA TCCGGGATCCCTGACCGATTCAAGTGGCAGCGGGTCTGGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCGATCACCTTTGGC CAAGGACACGACTGGAGATTAAA
31	CD33 AF5 HL x H2C HL	artificial	aa	QVQLVQSGAEVKKPGASVKVSCKASGYTFITNYGMNWVKQAPGQGLKWMGWINTYTGEPYADDFKG RVMTSDTSTSTAYLELHNLRSDDTAVYICARWSWSDGYVYFDYWGQGTITVTVSSGGGGSGGGGS GGGSDIVMTQSPDRLTVSLGERTTINCKSSQSVLDSSKNKNSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYICQQSAHFPITFGQGRLEIKSGGGSEVQILVESG GGLVQPGGSLKLSCAASGFTFNKYAMNWVRQAPGKLEWVARIRSKYNNYATYYADSVKDRFTISR DDSKNTAYLQMNLLKTEDTAVYICVRHGNFGNSYISYWAYWGQGTITVTVSSGGGGSGGGSGGGGS QTVVTTQEPRLTVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTFKFLAPGTPARFSG

[illegible]

[illegible]

				GATGATTCAAAAAACACTGCCCTATCTACAAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAATAGCTACATATCCTACTGGGCTTACTGGGGCCAAAGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGTCTGGCGGCGCGGCTCCGGTGGTGGTGTCT CAGACTGTTGTGACTCAGGAACCTTCACACCGTATCACCTGGTGGAAACAGTCACACTCACTTGT GGCTCCTCGACTGGGGCTGTTACATCTGGCAACTACCCAAACTGGGTCCAAACAAAAACCAGGTCAG GCACCCCGTGGTCTAATAGGTGGGACTAAGTTCCTCGCCCCGGTACTCCTGCCAGATTCTCAGGC TCCCTGCTTGGAGGCAAGGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAATAT TACTGTGTTCTATGTTACAGCAACCGCTGGGTGTTCGGTGGAGGAACCAAACTGACTGTCCCTA QVQLVQSGAEVKKPGEISVKVSKKASGYTFNRYGMNWKQAPGQGLKWMGNINITYTGEPTYADDFKG RVMTMTDTSTSTAYMEIRNLNRNDDTAVYYCARWSWSDDGYVYFDYWGQGTTVTVSS
37	CD33 VH of AC8	artificial	aa	NYGMN
38	CD33 HCDR1 of AC8	artificial	aa	WINTYTGEPTYADDFKG
39	CD33 HCDR2 of AC8	artificial	aa	WSWSDGYVYFDY
40	CD33 HCDR3 of AC8	artificial	aa	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGGTGGATAAACACCTACACTGGAGAGCCAAACATATGTGTGATGACTTCAAGGGA CGGTTACCATGACTACGGATACCTCTACCAGCACTGCCTATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGTGAGTGTGAGTTACTACGTTTACTTGAC TACTGGGGCCCAAGGCACCTACGGTCACCGTCTCCTCA
41	CD33 VH of AC8	artificial	nt	DIVMTQSPDLSLTIVSLGERTTINCKSSQSVLDSKKNKNSLAWYQQKPGQPPKILLISWASTRESGIPD RFGSGSGTDFTLTIDSLQPEDSATYYCQQSAHFPIITFGQGTREIK
42	CD33 VL of AC8	artificial	aa	KSSQSVLDSKKNKNSLA
43	CD33 LCDR1 of AC8	artificial	aa	WASTRES
44	CD33 LCDR2 of AC8	artificial	aa	QQSAHFPIIT
45	CD33 LCDR3 of AC8	artificial	aa	GACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAGAGGACCACCATCAAC TGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAGAAATAAGAACTCCTTAGCTTGGTACCAGCAG AAACCCAGGACAGCCCTCCTAAATTTACTCCTTTCCTGGGCATCTACGCGGGAATCCGGGATCCCTGAC CGATTTCAGTGGCAGCGGGTCTGGGACAGATTTTCACTCTCACTATTGACAGCCTGCAGCCTGAAGAT TCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCGATCACCTTTGGCCAAAGGACACGACTG
46	CD33 VL of AC8	artificial	nt	

47	CD33 HL of AC8	artificial	aa	GAGATTAAA QVQLVQSGAEVKKPGESVKVSKASGYTFTNYGMNVWKQAPQQLKWMGWINTYTGEPYADDFKG RVMTTDTSTSTAYMEIRNLRNDDTAVYYCARWSWDGYVYFDYWGQGTIVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSLDSSKNKNSLAWYQKPGQPPKLLLSWASTRE SGIPDRFSGSGSDTDFTLTIDSLQPEDSATYCYQSSAHFPIITFGQGTRELEIK CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAGCCTGGAGAGTCAGTCAAGGTCTCTCTGC AAGGCTAGCGGGTATACCTTCACAAACTATGGAATGAAGTGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGCTGGATAAACACCTACACTGGAGAGCAACATATGCTGATGACTTCAAGGGA CGGGTTACCATGACTACGGATACCTCTACAGCACTGCCTATATGAAATCCGAACTCAGAAAT GATGACACGGCTGTATATTAATGTGCGCGCTGGAGTTGGAGTGATGGTTACTACTGTTACTTTGAC TACTGGGGCCAAAGCACTACGGTACCGTCTCTCTCAGGTGGTGGTGGTCTTGGCGGCGGCTCC GGTGGTGGTGGTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGAGTGTGTCTCTGGCGGAG AGGACCACTCAACTGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAGAAATAAGAACTCCTTA GCTTGGTACCAGAGAAAACAGGACAGCTCTCTAAATTAATCTCTTCTGGGCACTACGCGGGAA TCCGGGATCCCTGACCGATTGAGTGGCAGCGGCTCTGGGACAGATTCTACTCTCACTATTGACAGC CTGCAGCCTGAAGATTCTGCAACTTACTATATTGTCAACAGTCTGCCCACCTTCCCGATCACTTTTGGC CAAGGACACGACTGGAGATTAAA
48	CD33 HL of AC8	artificial	nt	QVQLVQSGAEVKKPGESVKVSKASGYTFTNYGMNVWKQAPQQLKWMGWINTYTGEPYADDFKG RVMTTDTSTSTAYMEIRNLRNDDTAVYYCARWSWDGYVYFDYWGQGTIVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSLDSSKNKNSLAWYQKPGQPPKLLLSWASTRE SGIPDRFSGSGSDTDFTLTIDSLQPEDSATYCYQSSAHFPIITFGQGTRELEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNWRQAPGKGLEWVARIRSKYNNYATYYADSVKDRFTISR DDSKNTAYLQMNLLKTEDTAVYYCVRHGNFNSYISYWAYWGQGTIVTVSSGGGSGGGGSGGGGS QTVVTEPESLTVSPGGTVTLTTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAGCCTGGAGAGTCAGTCAAGGTCTCTCTGC AAGGCTAGCGGGTATACCTTCACAAACTATGGAATGAAGTGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGCTGGATAAACACCTACACTGGAGAGCAACATATGCTGATGACTTCAAGGGA CGGGTTACCATGACTACGGATACCTCTACAGCACTGCCTATATGAAATCCGCACTCAGAAAT GATGACACGGCTGTATATTAATGTGCGCGCTGGAGTTGGAGTGATGGTTACTACTGTTACTTTGAC TACTGGGGCCAAAGCACTACGGTACCGTCTCTCTCAGGTGGTGGTGGTCTTGGCGGCGGCTCC GGTGGTGGTGGTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCACTCAACTGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAGAAATAAGAACTCCTTA GCTTGGTACCAGAGAAAACAGGACAGCTCTCTAAATTAATCTCTTCTGGGCACTACGCGGGAA TCCGGGATCCCTGACCGATTGAGTGGCAGCGGCTCTGGGACAGATTCTACTCTCACTATTGACAGC CTGCAGCTTGAAGATTCTGCAACTTACTATATTGTCAACAGTCTGCCCACCTTCCCGATCACTTTTGGC CAAGGACACGACTGGAGATTAAAATCCGAGGTGGTGGCTCCGAGGTGCAGTGGTGCAGTCTGGA GGAGGATTGGTGCAGCTGGAGGGTCAATTGAAACTCTCATGTGCAGCCTCTGGATTCACTTCAAT AAGTACGCCATGAACCTGGGTCCGCCAGGCTCCAGGAAAGGGTTTGGAATGGGTGCTCGCATAGA
49	CD33 AC8 HL x H2C HL	artificial	aa	
50	CD33 AC8 HL x H2C HL	artificial	nt	

		artificial	aa	AGTAAATATAATTAATGCAACATATATATGCCGATTAGTGAAGACAGAGTTTACCATCTCCAGA GATGATTCAAAAAACACTGCCATATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGAACTTCGGTAATAGCTACATATCTTACTTGGGCTTACTGGGGCCAAAGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGTCTTGGCGCGCGGCTCCGGTGGTGGTGGTGTCT CAGACTGTGTGACTCAGGAACCTTCACCTACCGTATCACCATACTGGTGAACAGTACACTCACTTGT GGCTCCTCGACTGGGCTGTACATCTGGCTACTACCCAACTGGGTCCAACAAAAACAGGTCAG GCACCCCGTGGTCTAATAGGTGGACTAAGTTCTCGCCCCGGTACTCCTGCCAGATTCTCAGGC TCCCCGTCTGGAGGCAAGGCTGCCCTCACCCCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAATAT TACTGTGCTCTATGTTACAGCAACCGCTGGGTGTTCTGGTGGAGGAACCAAACTGACTGTCTCTA QVQLVQSGAEVKKPESVKVSKASGYTFNRYGMNWKQAPGQGLKWMGWINTYTGEPYADDFKG RVMTTDTSTSTAYMEIRNLRNDDTAVYYCARWSWSDGYVYFDYWGQGTFTVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSVLDSKKNKNSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGGTDFLTIDSLQPEDSATYYCQQSAHFPITFGQGRLEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAAAGFTFNSYAMNWRQAPGKGLEWVARIRSKYNNIYATYYADSVKGRFTISR DDSKNTAYLQMNNLKTEDTAVYYCVRHGNFNSYVSWWAYWGQGTILVTVSSGGGSGGGGSGGGGS QTVVTQEPISLTIVSPGGTIVTLITCGSSTGAVTSGNYPNWVQQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGGKAAITLISGVQPEDEAEFYCVLMYSNRWVFGGKTLTVL
52	CD33 AC8 HL x F12Q HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAATGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCAACATATGCTGATGACTTCAAGGGA CGGTTACCATGACTACGGATACCTCTACAGACTGCCATATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTAATGTGTGCGGCTGGAGTTGGAGTGAATGTTACTAGTTTACTTTGAC TACTGGGGCCAAAGCACTACGGTACCGTCTCCTCAGGTGGTGGTGTCTGGCGGCGGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAGAAATAAGAACTCCTTA GCTTGGTACCAGAGAAACAGGACAGCTCCTAAATTAATCTCTTCTTGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTGAGTGGCAGCGGCTTGGGACAGATTTCACCTTCCGATCACTTTGGC CTGCAGCCTGAAGATTCTGCAACTTACTATTTGTCAACAGTCTGCCACTTCCCGATCACTTTGGC CAAGGGACACGACTGGAGATTAAATCCGAGGTGGTGGCTCCGAGGTGAGTGGTGGTGGTGGTGGT GGAGGATTGGTGCAGCTGGAGGTTCAATTGAAATCTCATGTGCAGCTTGGATTCACCTTCAAT AGCTACGCCATGAACCTGGGTCCGCCAGGCTCCAGGAAAGGTTTGGAAATGGTTCGCTCGATAAGA AGTAAATATAATTAATGCAACATATATATGCGGATTAGTGAAGAGCAGGTTTACCATCTCCAGA GATGATTCAAAAAACACTGCCATATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGAACTTCGGTAATAGTACGTTTCTTGGTGGGCTTACTGGGGCCAAAGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGGTCTTGGCGGCGGGCTCCGGTGGTGGTGGTCT CAGACTGTGTGACTCAGGAACCTTCACCTACCGTATCACCATACTGGTGAACAGTACACTCACTTGT GGCTCCTCGACTGGGCTGTATACATCTGGCACTACCCAACTGGGTCCAACAAAAACAGGTCAG GCACCCCGTGGTCTAATAGGTGGGACTAAGTTCTCGCCCCGGTACTCCTGCCAGATTCTCAGGC TCCCCTGCTTGGAGGCAAGGCTGCCCTCACCCCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAATAT

53	CD33 AC8 HL x I2C HL	artificial	aa	TACTGTGTTTCTATGGTACAGCAACCGCTGGGTGTTTCGGTGGAGGAACCAAACTGACTGTCTCTA QVQLVQSGAEVKKPESVKVSCKASGYFTNYGMNWKQAPGQGLKWMGWINTYTGTEPTYADDFKG RVMTTSDTSTSTAYMEIRNLRNDDTAVYYCARWSWDGYVYFDYWGQGTFTVTVSSGGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSLDSSKNKNSLAWYQOKPGQPPLKLLISWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYCYQQSAHFPIITFGQGTRLFEEKSGGGSEVQLVESG GGILVQPGSLKLSCAAASGFTFNKYAMNWRQAPGKGLEWVARIRSKYNNIYATYADSVKDRFTISR DDSKNTAYLQMNLLKTEDTAVYYCVRHGNFGNSYISYWAYWGQGTILTVSSGGGGSGGGSGGGGS QTVVTQEPSTLTVSPGGTVTLTCGSTSTGAFTSGNYPNWQOKPGQAPRGLIGGTGFLAPGTPARFSG SLLGGKAALTLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL
54	CD33 AC8 HL x I2C HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGTATACCTTCACAAAATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGGTGGATAAAACACCTACACTGGAGAGCCAAATATATGCTGATGACTTCAAGGGA CGGTTACCATGACTACGGATACCTCTACCAGCACTGCCATATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTGAC TACTGGGGCCAAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTTCTTGGCGGCGCGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCAAGAATAAGAACTCCTTA GCTTGGTACCAGAGAAACAGGACAGCCTCCTAAATTTACTCCTTTCTCTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTCAGTGGCAGCGGTCTGGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCCTGAAGATTTGCAACTTACTATTTGTCAAACAGTCTGCCCACTTCCCGATCACTTTGGC CAAGGACACAGCTGGAGATTAAATCCGGAGGTGGTGGCTCCGAGGTGCAGTGGTTCGAGTCTGGA GGAGGATTGGTGCAGCTGGAGGGTCAATTGAAATCTCTATGTGCAGCCTCTTGGAATTCACCTTCAAT AAGTACGCCATGAACTGGGTCCCGCAGGCTCCAGGAAAGGGTTTGGAAATGGGTGCTCGCATAGA AGTAAATATAATTAATATGCAACATATTTATGCCGATTTCAGTGAAAGACAGGTTACCATCTCCAGA GATGATTCAAAAAACACTGCCATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAAATAGCTACATATCCTTACTTGGGCTTACTTGGGGCCAAAGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGGTCTTGGCGGCGCGGCTCCGCTGGTGGTGGTCTT CAGACTGTTGTGACTCAGGAACCTTCACCTACCGTATCACCTGGTGGAACAGTCACACTCACTGT GGCTCCTCGACTGGGGCTGTTACATCTGGCAACTACCCAACTGGGTCCAAACAAAAACAGGTCAG GCACCCCGTGGTCTAATAGGTGGGACTAAAGTTCCCTCGCCCCGGTACTCCTGCCAGATTTCCAGGC TCCCTGCTTGGAGGCAAGGCTGCCCTCACCCCTCTCAGGGTGACAGCCAGAGGATGAGGCAGAAATAT TACTGTGTTCTATGTTACAGCAACCGCTGGGTGTTTCGGTGGAGGAACCAAACTGACTGTCTCTA QVQLVQSGAEVKKPESVKVSCKASGYFTNYGMNWKQAPGQGLKWMGWINTYTGTEPTYADDFKG RVMTTSDTSTSTAYMEIRSLRSDDTAVYYCARWSWDGYVYFDYWGQGTFTVTVSS NYGMN
55	CD33 VH of AH11	artificial	aa	
56	CD33 HCDR1 of AH11	artificial	aa	
57	CD33 HCDR2 of AH11	artificial	aa	WINTYTGTEPTYADDFKG

58	HCDR3	artificial	aa	WSWSDGYVYVFDY	
59	CD33 VH of AH11	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCAACATATGCTGATGACTTCAAGGGA CGGGTTACCATGACTTCGGATACCTCTACACAGCACTGCCCTATATGGAATTCAGCAGCCTCAGAAAT GATGACACGGCTGTATATTAATGCTGTGCGCTGGAGTTGGAGTATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGCACTACGGTCACCGTCTCCTCA	
60	CD33 VL of AH11	artificial	aa	DIVMTQSPDSLTVSLGERTTINCKSSQSVLDSSKNKNSLAWYQQKPGQPPLKLLLSWASTRESGIPD RFSGSGGTDFLTITIDSLQPEDSATYYCQSAHFPIITFGQGTRLEIK	
61	CD33 LCDR1 of AH11	artificial	aa	KSSQSVLDSSKNKNSLA	
62	CD33 LCDR2 of AH11	artificial	aa	WASTRES	
63	CD33LCDR3 of AH11	artificial	aa	QQSAHFPIIT	
64	CD33 VL of AH11	artificial	nt	GACATCGTGATGACACAGCTCTCCAGACTCCCTGACTGTGTCTCTGGGCGAGAGGACCACCATCAAC TGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAGAAATAAGAACTCCTTAGCTTGGTACCAGCAG AAACCAGGACAGCCTCCTAAATTAATCTCTTCTTGGGCATCTACGCGGGAATCCGGGATCCCTTGAC CGATTGAGTGGCAGCGGGTCTGGGACAGATTTCATCTCACTATTGACAGCCTGCAGCCTGAAGAT TCTGCAACTTACTATTGTCAACAGCTCTGCCACTTCCCGATCACCTTTGGCCAAAGGACACAGACTG GAGATTAAA	
65	CD33 HL of AH11	artificial	aa	QVQLVQSGAEVKKPGEISVKVSKASGYTFITNYGMNWVKQAPGQGLKMMGWINTYTGEPYADDFKG RVTMTSDTSTSTAYMEISSLRSDDTAVYICARWSWSDGYVYFDYWGQGTFTVTVSSGGSGGGGS GGGSDDIVMTQSPDSLTVSLGERTTINCKSSQSVLDSSKNKNSLAWYQQKPGQPPLKLLLSWASTRE SGIPDRFSGSGGTDFLTITIDSLQPEDSATYYCQSAHFPIITFGQGTRLEIK	
66	CD33 HL of AH11	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGCTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCAACATATGCTGATGACTTCAAGGGA CGGGTTACCATGACTTCGGATACCTCTACACAGCACTGCCCTATATGGAATTCAGCAGCCTCAGAAAT GATGACACGGCTGTATATTAATGCTGTGCGCTGGAGTTGGAGTATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGCACTACGGTCACCGTCTCCTCAGGTGGTGGTGTCTGGCGGCGGCTCC GGTGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAGAAATAAGAACTCCCTTA GCTTGGTACCAGAGAAAACAGGACAGCCTCCTAAATTAATCTCTTCTGGGCATCTACGCGGGA TCCGGGATCCCTTGACCGGATTCAGTGGCAGCGGTCTGGGACAGATTTCATCTTCATATTGACAGC CTGCAGCCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCGATCACCTTTGGC	

67	CD33 AH11 HL x H2C HL	artificial	aa	CAAGGACACGACTGGAGATTAAA QVQLVQSGAEVKKPESVKVSCKASGYTFITNYGMNWVKQAPGQGLKWMGWINTYITGEPTYADDFKG RVMTSDTSTSTAYMEISSLRSDDTAVYYCARWSWDGYVYFDYWGQGTITVTVSSGGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSLDSSKNKNSLAWYQOKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYYCQOQSAHFPIITFGQGTREIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAAASGFTFNKYAMNWRQAPGKGLEWVARIRSKYNNIYATYYADSVKGRFTISR DDSKNTAYLQMNLIKTEDTAVYYCVRHGNFGNSIYSWAYWGQGTITVTVSSGGGGSGGGSGGGGS QTVVTQEPISLTVSPGGTVTLITCGSSSTGAVTSGYKPNWVQOKPGQAPRGLIGGTKFLAPGTPARFSG SLLGGKAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLITVL
68	CD33 AH11 HL x H2C HL	artificial	nt	CAGGTGCAGCTGGTGACGTGGAGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGTC AAGGCTAGCGGGTATACCTTCACAAAACATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGGTGGATAAAACACCTACACTGGAGAGCCAAACATATGCTGATGACTTCAAGGGA CGGTTACCATGACTTCGGATACCTCTACCAGCACTGCCATATATGGAATTCAGCAGCCTCAGAAAT GATGACACGGCTGTATATTTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTTCTTGGCGGCGCGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCAAGAATAAGAACTCCTTA GCTTGGTACCAGAGAAAACAGGACAGCCTCCTAAATTTACTCCTTTCTCTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGGATTCAGTGGCAGCGGTCTGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCCTGAAGATTCGCAACTTACTATTTGTCACACAGTCTGCCACTTCCCGATCACTCTTGGC CAAGGACACGACTGGAGATTAAATCCGGAGGTGGTGGCTCCGAGGTGCAGCTGGTCGAGTCTGGA GGAGGATTGGTGCAGCTGGAGGGTCAATTGAAACTCTCATGTGCAGCCTCTGGATTCACTTCAAT AAGTACGCCATGAACTGGGTCCCGCAGGCTCCAGGAAAGGGTTTGGAAATGGGTGCTCGCATAGA AGTAAATATAATTAATATGCAACATATTTATGCCGATTAGTGAAGACAGGTTACCATCTCCAGA GATGATTCAAAAAACACTGCCATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAAATAGCTACATATCCTTACTTGGGCTTACTTGGGGCCAAAGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGTCTTGGCGGCGCGGCTCCGGTGGTGGTGGTCTT CAGACTGTTGTGACTCAGGAACCTTCACTACCGTATCACTGGTGGAACAGTCACACTCACTGT GGCTCCTCGACTGGGGCTGTTACATCTGGCTACTACCCAAACTGGGTCCAAACAAAAACAGGTCAG GCACCCCGTGGTCTAATAGGTGGACTAAAGTTCTTCCCGCGGTACTCCTTGCAGATTCTCAGGC TCCCTGTCTTGGAGGCAAGGCTGCCCTCACTCCCTCTCAGGGTCAAGCCAGAGGATGAGGCAGAAATAT TACTGTGCTCTATGTTACAGCAACCGCTGGGTGTTCTGGTGGAGGAAACCAAACTGACTGTCTCTA
69	CD33 AH11 HL x F12Q HL	artificial	aa	QVQLVQSGAEVKKPESVKVSCKASGYTFITNYGMNWVKQAPGQGLKWMGWINTYITGEPTYADDFKG RVMTSDTSTSTAYMEISSLRSDDTAVYYCARWSWDGYVYFDYWGQGTITVTVSSGGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSLDSSKNKNSLAWYQOKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYYCQOQSAHFPIITFGQGTREIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAAASGFTFNKYAMNWRQAPGKGLEWVARIRSKYNNIYATYYADSVKGRFTISR DDSKNTAYLQMNLIKTEDTAVYYCVRHGNFGNSIYSWAYWGQGTITVTVSSGGGGSGGGSGGGGS QTVVTQEPISLTVSPGGTVTLITCGSSSTGAVTSGNYPNWVQOKPGQAPRGLIGGTKFLAPGTPARFSG

70	CD33 AH11 HL x F12Q HL	artificial	nt	SLGGKAALTL.SGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAACCTGGAGAGTCAGTCAAGGTCTCCCTGC AAGGCTAGCGGGTATACCTTCACAAAACATGGAATGAAC'TGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCCAACATATGCTGATGACTTCAAGGGA CGGGTTACCATGACTTCGGATACCTCTACCAGCACTGCC'TATATGGAATCAGCAGCCTCAGAAAT GATGACACGGCTGTATATTA'CTGTGCGCGTGGAGTTGGAGTGATGGTTACTACGTTTACTTTTGAC TACTGGGGCCAAGGCACTACGGTCACCGTCTCCTCAGGTGGTGGTGGTTCTTGGCGGGCGGCTCC GGTGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGT'TTAGACAGTCCAAGAA'TAAGAACTCCTTTA GCTTGGTACCAGAGAAACCAGGACAGCCTCCTAAATTA'CTCTTCTTGGGCATCTACGCGGGA TCCGGGATCCCTGACCGATT'CA'GTGGCAGCGGTCTGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCC'GATCACCTTTGGC CAAGGGACAGCTGGAGATTAAATCCGAGGTGGTGGTCCGAGGTGCAGTGGTGCAGTCTGGA GGAGGATTGGTGCAGCTGGAGGTCA'TTGAACCTCTCATGTGCAGCCTCTGGATTCACTTCAAT AGCTACGCCATGAAC'TGGTCCGCCAGGCTCCAGGAAAGGTTTGGAA'TGGTGTCTCGCATAAAGA AGTAAATATAATAATTA'GTCAACATATTATGCCGATT'CA'GTGAAAGGCAGGTTCA'CCATCTCCAGA GATGATTCAAAAAACACTGCC'TATCTACAAATGAACAACTTGA'AAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAAATAGCTACGTTTCTTGGTGGGTACTTGGGGCCAAAGGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGGTTC'TGGCGCGCGGCTCCGGTGGTGGTGGTTCCT CAGACTGTGTGACTCAGGAACTTCACTCA'CCGTATCACTGGTGGAA'CACTCACTCACTTGT GGCTCTCGACTGGGCTGT'TACATCTGGCAACTACCCAACTGGGTCCAACAA'AACTCAGGTCA'G GCACCCGTGGTCTAAATAGTGGGACTAAGTTCTCTGCCCCCGGTACTCTGCCAGATTCTCAGGC TCCCTGCTTGGAGGCAAGGTGCCCTCACCTCTCA'GGGTACAGCCAGAGGATGAGGCAAGAA'TAT TACTGTGTCTATGTTACAGCAACCGTGGGTGTTTCGGTGGAGGAA'CAAACTGACTGTCTCTA
71	CD33 AH11 HL x I2C HL	artificial	aa	QVQLVQSGAEVKKPESVKVSCKASGYTFFTNYGMNWVKQAPGQGLKMWGINTYITGEPTYADDFKG RV'TMTSDTSTAYMEISSLRSDDTAVYYCARWSWSDGYVYFDYWQGTTVTVSSGGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSVLDSKNKNSLAWYQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYYCQSAHFPIITFGQGTRELEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNW/RQAPGKGLEWVARIRSKYNNYATYYADSVKDRFTIISR DDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYISYWAYWQGTGLTVTVSSGGGGSGGGSGGGGS QTVV'TQEP'SLTVSPGGTVTLTCGSSTGA'VTS'GNYPNWVQKPGQAPRGLIGGTFKFLAPGTPARFSG SLGGKAALTL.SGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL CAGGTGCAGCTGGTGCAGCTTCGGAGCTGAGGTGAAGAACCTTCGGAGAGTCAGTCAAGGTCTCCCTGC AAGGCTAGCGGGTATACCTTCACAAAACATGGAATGAAC'TGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCCAACATATGCTGATGACTTCAAGGGA CGGGTTACCATGACTTCGGATACCTCTACCAGCACTGCC'TATATGGAATCAGCAGCCTCAGAAAT GATGACACGGCTGTATATTA'CTGTGCGCGTGGAGTTGGAGTGATGGTTACTACGTTTACTTTTGAC TACTGGGGCCAAGGCACTACGGTCACCGTCTCCTCAGGTGGTGGTGGTTCTTGGCGGGCGGCTCC GGTGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGT'TTAGACAGTCCAAGAA'TAAGAACTCCTTTA GCTTGGTACCAGAGAAACCAGGACAGCCTCCTAAATTA'CTCTTCTTGGGCATCTACGCGGGA TCCGGGATCCCTGACCGATT'CA'GTGGCAGCGGTCTGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCC'GATCACCTTTGGC CAAGGGACAGCTGGAGATTAAATCCGAGGTGGTGGTCCGAGGTGCAGTGGTGCAGTCTGGA GGAGGATTGGTGCAGCTGGAGGTCA'TTGAACCTCTCATGTGCAGCCTCTGGATTCACTTCAAT AGCTACGCCATGAAC'TGGTCCGCCAGGCTCCAGGAAAGGTTTGGAA'TGGTGTCTCGCATAAAGA AGTAAATATAATAATTA'GTCAACATATTATGCCGATT'CA'GTGAAAGGCAGGTTCA'CCATCTCCAGA GATGATTCAAAAAACACTGCC'TATCTACAAATGAACAACTTGA'AAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAAATAGCTACGTTTCTTGGTGGGTACTTGGGGCCAAAGGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGGTTC'TGGCGCGCGGCTCCGGTGGTGGTGGTTCCT CAGACTGTGTGACTCAGGAACTTCACTCA'CCGTATCACTGGTGGAA'CACTCACTCACTTGT GGCTCTCGACTGGGCTGT'TACATCTGGCAACTACCCAACTGGGTCCAACAA'AACTCAGGTCA'G GCACCCGTGGTCTAAATAGTGGGACTAAGTTCTCTGCCCCCGGTACTCTGCCAGATTCTCAGGC TCCCTGCTTGGAGGCAAGGTGCCCTCACCTCTCA'GGGTACAGCCAGAGGATGAGGCAAGAA'TAT TACTGTGTCTATGTTACAGCAACCGTGGGTGTTTCGGTGGAGGAA'CAAACTGACTGTCTCTA
72	CD33 AH11 HL x I2C HL	artificial	nt	QVQLVQSGAEVKKPESVKVSCKASGYTFFTNYGMNWVKQAPGQGLKMWGINTYITGEPTYADDFKG RV'TMTSDTSTAYMEISSLRSDDTAVYYCARWSWSDGYVYFDYWQGTTVTVSSGGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSVLDSKNKNSLAWYQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYYCQSAHFPIITFGQGTRELEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNW/RQAPGKGLEWVARIRSKYNNYATYYADSVKDRFTIISR DDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYISYWAYWQGTGLTVTVSSGGGGSGGGSGGGGS QTVV'TQEP'SLTVSPGGTVTLTCGSSTGA'VTS'GNYPNWVQKPGQAPRGLIGGTFKFLAPGTPARFSG SLGGKAALTL.SGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL CAGGTGCAGCTGGTGCAGCTTCGGAGCTGAGGTGAAGAACCTTCGGAGAGTCAGTCAAGGTCTCCCTGC AAGGCTAGCGGGTATACCTTCACAAAACATGGAATGAAC'TGGGTGAAGCAGGCTCCAGGACAGGGT TTAAAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCCAACATATGCTGATGACTTCAAGGGA CGGGTTACCATGACTTCGGATACCTCTACCAGCACTGCC'TATATGGAATCAGCAGCCTCAGAAAT GATGACACGGCTGTATATTA'CTGTGCGCGTGGAGTTGGAGTGATGGTTACTACGTTTACTTTTGAC TACTGGGGCCAAGGCACTACGGTCACCGTCTCCTCAGGTGGTGGTGGTTCTTGGCGGGCGGCTCC GGTGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGT'TTAGACAGTCCAAGAA'TAAGAACTCCTTTA GCTTGGTACCAGAGAAACCAGGACAGCCTCCTAAATTA'CTCTTCTTGGGCATCTACGCGGGA TCCGGGATCCCTGACCGATT'CA'GTGGCAGCGGTCTGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCC'GATCACCTTTGGC CAAGGGACAGCTGGAGATTAAATCCGAGGTGGTGGTCCGAGGTGCAGTGGTGCAGTCTGGA GGAGGATTGGTGCAGCTGGAGGTCA'TTGAACCTCTCATGTGCAGCCTCTGGATTCACTTCAAT AGCTACGCCATGAAC'TGGTCCGCCAGGCTCCAGGAAAGGTTTGGAA'TGGTGTCTCGCATAAAGA AGTAAATATAATAATTA'GTCAACATATTATGCCGATT'CA'GTGAAAGGCAGGTTCA'CCATCTCCAGA GATGATTCAAAAAACACTGCC'TATCTACAAATGAACAACTTGA'AAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAAATAGCTACGTTTCTTGGTGGGTACTTGGGGCCAAAGGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGGTTC'TGGCGCGCGGCTCCGGTGGTGGTGGTTCCT CAGACTGTGTGACTCAGGAACTTCACTCA'CCGTATCACTGGTGGAA'CACTCACTCACTTGT GGCTCTCGACTGGGCTGT'TACATCTGGCAACTACCCAACTGGGTCCAACAA'AACTCAGGTCA'G GCACCCGTGGTCTAAATAGTGGGACTAAGTTCTCTGCCCCCGGTACTCTGCCAGATTCTCAGGC TCCCTGCTTGGAGGCAAGGTGCCCTCACCTCTCA'GGGTACAGCCAGAGGATGAGGCAAGAA'TAT TACTGTGTCTATGTTACAGCAACCGTGGGTGTTTCGGTGGAGGAA'CAAACTGACTGTCTCTA

				AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTTTTAGACAGCTCCAAGAAATAAGAACTCCCTTA GCTTGGTACCAGCAGAAAAACAGGACAGCCTCCATAATTACTCCTTCCCTGGGCATCTACGCGGAAA TCCGGGATCCCTGACCGATTCAAGTGGCAGCGGGTCTGGGACAGATTTCACTCTCACTATTTGACAGC CTGAGCCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCGATCACCTTTGGC CAAGGACACGACTGGAGATTAAATCCGAGGTGGTGGTCCGAGGTGAGTGGTCGAGTCTGGA GGAGGATTGGTGACGCTGGAGGTCATTGAAACTCTCATGTGCAGCCTCTGGATTCACTTCAAT AAGTACGCCATGAACTGGGTCGCCAGGCTCCAGGAAAGGTTTGAAATGGGTGCTCGCATAAAGA AGTAAATATAATAATTATGCAACATATTATGCCGATTCAAGTGAAGACAGGTTCAACATCTCCAGA GATGATTCAAAAACACTGCCCTATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGAACTTCGGTAATAGCTACATATCTACTGGCTTACTGGGCCAAGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGTCTGGCGCGCGGCTCCGGTGGTGGTGGTCT CAGACTGTTGTGACTCAGGAACCTTCACTCACCGTATCACTGGTGGAAACAGTCACACTCACTTGT GGCTCCTCGACTGGGCTGTTACATCTGGCAACTACCCAACTGGGTCCAACAAAAACAGGTCAG GCACCCCGTGGTCTAAATAGGTGGACTAAGTTCTCGCCCCCGGTACTCTCTGCCAGATTCTCAGGC TCCCCTGCTTGGAGGCAAGGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGTTCTATGTTACAGCAACCGCTGGGTGTTCTGGTGGAGGAAACCAAACTGACTGTCTCTA
73	CD33 VH of B3	artificial	aa	QVQLVQSGAEVKKPESVKVSCKASGYTFNTNYGMNWKQAPQGLEWMGWINTYTGENTNYADKFQG RVFTSTSTSTAYMEILRNLIKSDDTAVYYCARWSWSDGYVYFDYWGQGTTVTTVSS
74	CD33 HCDR1 of B3	artificial	aa	NYGMN
75	CD33 HCDR2 of B3	artificial	aa	WINTYTGENTNYADKFQG
76	CD33 HCDR3 of B3	artificial	aa	WSWSDGYVYFDY
77	CD33 VH of B3	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGTC AAGGCTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGGTGGATAAACACCTACACTGGAGAGACAAACTATGCTGATAAGTTCCAGGGA CGGTTACCTTCACTTCGGATACCTCTACCAGCACTGCCATATATGGAACCTCCGCAACCTCAAAAAGT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGTTACTACGTTTACTTTTGAC TACTGGGGCCAAAGGCACATACGGTCACCGTCTCCTCA
78	CD33 VL of B3	artificial	aa	DIVMTQSPDSMTVSLGERTTINCKSSQSVLDSSTNKNLSLAWYQQKPGQPPKLLLSWASTRESGIPD RFSGSGSGTDFTLTIDSLQPEDSATYCYQQSAHFPITFTFGQGTRLDIK
79	CD33 LCDR1 of B3	artificial	aa	KSSQSVLDSSTNKNLSLA
80	CD33 LCDR2 of B3	artificial	aa	WASTRE
81	CD33LCDR3 of B3	artificial	aa	QQSAHFPIT
82	CD33 VL of B3	artificial	nt	GACATCGTGATGACACAGTCTCCAGACTCCATGACTGTGTCTCTGGGCGAGAGGACCACCATCAAC TGCAAGTCCAGCCAGAGTGTTTTAGACAGCTCCACGAAATAAGAACTCTTACTTGGTACCAGCAG

83	CD33 HL of B3	artificial	aa	AAAACAGGACAGCCTCCTAAATTTACTCCTTTCTTGGGCACTTACGCGGGAAATCCGGGATCCCTTGAC CGATTCAGTGGCAGCGGGTCTGGGACAGATTTCACTCTCACTATTGACAGCCTGCAGCCTGAAGAT TCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCGATCACCTTTGGCCAAAGGACACGACTG GACATTAAA QVQLVQSGAEVKKPGEISVKVSKKASGYTFITNYGMNWVKQAPGQGLEWMGWINTYTGFTNYADKFKQG RVFTFTSDTSTAYMELRNLIKSDDTAVYYCARWSWSDGYVYFDYWQGTITVTVSSGGGSGGGGS GGGSDIVMTQSPDSMTVSLGERITINCKSSQSVLDSSSTNKNLSLAWYQQKPGQPPKILLLSWASTRE SGIPDRFSGSGSDFTLTIDSLQPEDSATYFCQQSAHFPITFGQGTSLDIK
84	CD33 HL of B3	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAAGTCAAGGTCTCCTGTC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGCT TTAGAGTGGATGGGCTGGATAAAACACCTACACTGGAGAGACAAAATATGCTGATAAAGTTCCAGGGA CGGTTACCTTCACTTCGGATACCTTACAGCAGCTGCTTATATGGAACCTCCGCAACCTCAAAAGT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGTGTTACTACGTTACTTTGAC TACTGGGCGCAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTCTTGGCGGCGGGCTCC GGTGGTGGTGGTCTTGACATCGTATGACACAGCTCCAGACTCCATGACTGTGCTCTTGGGCGAG AGGACCACTCACTGCAAGTCCAGCCAGAGTGTTTTGAACAGCTCCAGCAATAAGAACTCCTTA GCTTGGTACCAGAGAAAACAGGACAGCCTCCTAAATTTACTCCTTTCCCTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTGAGTGCAGTGCAGCGGCTCTGGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCGATCACCTTTGGC CAAGGACACGACTGGACATTAAA
85	CD33 B3 HL x H2C HL	artificial	aa	QVQLVQSGAEVKKPGEISVKVSKKASGYTFITNYGMNWVKQAPGQGLEWMGWINTYTGFTNYADKFKQG RVFTFTSDTSTAYMELRNLIKSDDTAVYYCARWSWSDGYVYFDYWQGTITVTVSSGGGSGGGGS GGGSDIVMTQSPDSMTVSLGERITINCKSSQSVLDSSSTNKNLSLAWYQQKPGQPPKILLLSWASTRE SGIPDRFSGSGSDFTLTIDSLQPEDSATYFCQQSAHFPITFGQGTSLDIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAAAGTTFNKYAMNWVRQAPGKLEWVARIRSKYNNYATYYADSVKDRFTISR DDSKNTAYLQMNLIKTEDTAVYYCVRHGNTSYISYWAYWQGTITVTVSSGGGSGGGGSGGGGS QTVVTFEPSTLVSPGGTITLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGGKAALTLISGVQPEDEAEFYCALWYSNRWVFGGGTKLTVL
86	CD33 B3 HL x H2C HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAAGTCAAGGTCTCCTGTC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGCT TTAGAGTGGATGGGCTGGATAAAACACCTACACTGGAGAGACAAAATATGCTGATAAAGTTCCAGGGA CGGTTACCTTCACTTCGGATACCTTACAGCAGCTGCTTATATGGAACCTCCGCAACCTCAAAAGT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGTGTTACTACGTTACTTTGAC TACTGGGCGCAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTCTTGGCGGCGGGCTCC GGTGGTGGTGGTCTTGACATCGTATGACACAGCTCCAGACTCCATGACTGTGCTCTTGGGCGAG AGGACCACTCACTGCAAGTCCAGCCAGAGTGTTTTGAACAGCTCCAGCAATAAGAACTCCTTA GCTTGGTACCAGAGAAAACAGGACAGCCTCCTAAATTTACTCCTTTCCCTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTGAGTGCAGTGCAGCGGCTCTGGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCGATCACCTTTGGC

				CAAGGGACACGACTGGACATTTAAATCCGGAGGTGGTGGCTCCGAGGTGCAGTGGTTCGAGTCTGGA GGAGGATTGGTGCAGCCTGGAGGGTCAATTGAAACTCTCATGTGCAGCCTCTGGATTCACTTCAAT AAGTACGCCATGAACCTGGGTCCGCCAGGCTCCAGGAAAGGGTTTGGAAATGGGTTCGTCGCATAAGA AGTAAATATAAATAATATGCAACATATATATGCCGATTAGTGAAGACAGAGTTTACCATCTCCAGA GATGATTCAAAAAACACTGCCTATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAATAGCTACATATCTTCTGCGCGCGGCTCCGGTGGTGGTTCT ACTCTGGTCAACCGTCTCCTCAGGTGGTGGTGGTGGTGGCGCGGCTCCGGTGGTGGTGGTGGTGGT CAGACTGTGTGACTCAGGAACTTCACCTACCGTATCACCTTGGTGGAAACAGTACACTCACCTTGT GGCTCCTCGACTGGGGCTGTACATCTGGCTACTCCCAAACTGGGTCCAACAAAAACAGGTTCAG GCACCCCGTGGTCTAAATAGGTGGGACTAAGTCTCTCCCCCGGTACTCTCTGCCAGATCTCTCAGGC TCCCCTGCTTGGAGGCAAGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGCTCTATGGTACAGCAACCGCTGGGTGTTCCGTGGAGGAAACCAAACTGACTGTCTCTA
87	CD33 B3 HL x F12Q HL	artificial	aa	QVQLVQSGAEVKKPESVKVSCKASGYTFNMGMMWVKQAPQQLLEWMGWINTYTGENTYADKFQ RVFTTSDTSTAYMELRNILKSDDTAVYYCARWSWSGDYVYFDYWGQGTTVTVSSGGGGSGGGGS GGGSDIVMTQSPDSMTVSLGERTTINCKSSQSVLDSSTNKNSLAWYQKPGQPPKLLLSWASTRE SGIPDRFSGSGSDTDFTLTIDSLQPEDSATYCYQQSAHFPITFTGQGTSLDIKSGGGGSEVQLVESG GGIVQPGGSLKLSCAASGFTFNSYAMNWRQAPGKGLEWVARIRSKYNNIYATYADSVKGRFTISR DDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYVSWWAYWGQTLTVTVSSGGGGSGGGSGGGGS QTVVTFEPLSLTVSPGGTVTLTCGSSTGAVTSGNYPNWVQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEYYCVLMWSNRWVFGGTFKLTIVL
88	CD33 B3 HL x F12Q HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGCTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGGTGGATAAACACCTACACTGGAGAGACAAACTATGCTGATAAGTTCCAGGGA CGGTTACCTTCACCTCGGATACCTCTACCAGCACTGCCATATATGGAACCTCCGCAACCTCAAAAGT GATGACACGGCTGTATATTTACTGTGCGCGCTGGAGTTGGAGTGAGTTACTACGTTTACTTTTGAC TACTGGGGCCAAAGGCATACGGTCAACCGTCTCCTCAGGTGGTGGTGGTGGTGGTGGTGGTGGTGGT GGT AGGACCAACCATCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCACGAATAAGAACTCCTTA GCTTGGTACCAGCAGAAACCCAGGACAGCCTCCTAAATTTACTCCTTTCTTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGGATTCAGTGGCAGCGGTCTGGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCCTGAAAGATTCTGCAACTTACTATTTGTCAACAGTCTGCCCACTTCCCGGATCACCTTTGGC CAAGGGACACGACTGGACATTTAAATCCGGAGGTGGTGGTCCGAGGTGCAGTGGTTCGAGTCTGGA GGAGGATTGGTGCAGCTGGAGGGTCAATTGAAACTCTCATGTGCAGCCTCTGGAATCACTTCAAT AGTACGCCATGAACCTGGGTCCCGCAGGCTCCAGGAAAGGGTTTGGAAATGGGTTCGTCGCATAAGA AGTAAATATAAATAATATGCAACATATTTATGCCGATTAGTGAAGAGCAGGTTTACCATCTCCAGA GATGATTCAAAAAACACTGCCATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAATAGCTACGTTTCTTGGTGGGCTTACTGGGGCCAAAGG ACTCTGGTCAACCGTCTCCTCAGGTGGTGGTGGTGGTGGTGGTGGTGGTGGTGGTGGTGGTGGTGGT CAGACTGTGTGACTCAGGAACTTCACCTACCGTATCACCTGGTGGAAACAGTACACTCACTTGT

					GGCTCCCTCGACTGGGCTGTTACATCTGGCAACTTACCCTGGGTCCCAACAAAAACCCAGGTCAGGCACCCCGTGGTCTAATAGGTGGGACTAAGTTCTCGCCCCCGGTACTCTCGCCAGATTTCTCAGGCTCCCTGCTTGAGGCAAGGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGATGAGGCAGAAATATTACTGTGTTCTATGGTACAGCAACCGCTGGGTGTTCCGTGGAGGAACCAAACTGACTGTCTCCTA
89	CD33 B3 HL x I2C HL	artificial	aa		QVQLVQSGAEVKKPESVKVSCKASGYTFTNYGMNWVKQAPGQGLEWMGWINTYTGENTYADKFKQGRTVFTSDTSTSTAYMELRNILKSDDTAVYYCARWSWSDGYVYFDYWQGTGTTVTVSSGGGGGGGGSGGGSDIVMTQSPDSMTVSLGERTTINCKSSQSVLDSSTNKNISLAWYQKPGQPPKLLLSWASTRESGIPDRFSGSGGTDFTLTIDSLQPEDSATYYCQSAHFPIITFGQGTRLDIKSGGGSEVQLVESGGGLVQPGGSLKLSCAASGFTFNKYAMNW/RQAPGKGLEWVARIRSKYNNYATYYADSVKDRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNFGNSYISYWAYWGQGTIVTVSSGGGGSGGGSGGGGSQTVVTQEPSTLTVSPGGTVTLTCGSSTGAVTSGNYPNWVQKPGQAPRGLIGGTKFLAPGTPARFSGSLGGKAAALTLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLTIVL
90	CD33 B3 HL x I2C HL	artificial	nt		CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAGCCTGGAGAGTCAGTCAAGGTCTCCTGCAAGGTAGCGGGTATACCTTCACAAACTATGGAATGAACCTGGGTGAAGCAGGTCACAGCACAGGTTTAGAGTGGATGGGTGGATAAACACCTACACTGGAGAGACAAACTATGCTGATAAGTTCCAGGATCGGTTACCTTCACTTCGGATACCTCTACACGACCTGCCTATATGGAACCTCCGCAACCTCAAAAGTGATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTATGTTACTACGTTTATGACCTTACTGGGGCCAAAGCACACGGTCACCGTCTCCTCAGGTGGTGGTGGTCTGGCGGCGCGGCTCCGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCATGACTGTGTCTCTGGGCGAGAGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTTTTAGACAGCTCCACGAATAAGAACTCCTTATGCTTGTTACAGCAGAAAAACAGGACAGCCTCCTAAATTACTCCTTTCTGGGCATCTACGCGGAAATCCGGGATCCTTGACCGATTCAGTGGCAGCGGGTCTGGGACAGATTTCACTCTCACTATTGACAGCTGCAGCCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCAGTACACTTTGGCCAAGGACACGACTGGACATTAAATCCGGAGGTGGTGGTCCGAGGTGACGTGGTCGAGTCTGGAAGGATTGGTGACGCTGGAGGTCATTGAAACTCTCATGTGCAGCCTCTGGATTCACTTCAATCAATAGTACGCCATGAACGGTCCGCCAGGCTCCAGGAAAGGTTTGAAATGGGTGTCGCGATAAGAAGTAAATATAATAATTATGCAACATATTATGCCGATTCAGTGAAAAGACAGGTTTACCATTCTCCAGATGATTCAAAAAACACTGCCATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTACTGTGTGAGACATGGAACTTCGGTAATAGTACATATCCTACTGGCTTACTTGGGCGCAAGGACCTTGCTGGTCACCGTCTCCTCAGGTGGTGGTTCCTGGCGCGGCTCCCGTGGTGGTTCCTCAGACTGTGTGACTCAGGAACCTTCACTCACCGTATCACTGGTGGAAACAGTCACTCACTTGTGGCTCTCGACTGGGCTGTACATCTGGCAACTACCCAACTGGGTCCAAACAAACCCAGGTCAGGCACCCCGTGGTCTAATAGGTGGGACTAAGTTCTCGCCCCCGTACTCTGCCAGATTCTCAGGCTCCCTGCTTGAGGCAAGGTGCCCTCACCTCTCAGGGGTACGCCAGAGATGAGGCAGAAATATTACTGTGTTCTATGTTACAGCAACCGCTGGGTGTTCCGTGGAGGAACCAAACTGACTGTCTCCTA
91	CD33 VH of F2	artificial	aa		QVQLVQSGAEVKKPESVKVSCKASGYTFTNYGMNWVKQAPGQGLEWMGWINTYTGENTYADKFKQGRVFTSDTSTSTAYMELRNILKSDDTAVYYCARWSWSDGYVYFDYWQGTGTTVTVSS
92	CD33 HCDR1 of F2	artificial	aa		NYGMN

93	CD33 HCDR2 of F2	artificial	aa	WINTYTGETNYADKFGQ
94	CD33 HCDR3 of F2	artificial	aa	WSWSDGYVYVFDY
95	CD33 VH of F2	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAACACCTACACTGGAGAGACAAAATATGCTGATAAGTTCCAGGGA CGGTTACCTTCACCTTCGATACCTCTACCAGCACTGCCTATATGGAACCTCCGCAACCTCAAAAGT GATGACACGGCTGTATATTAATGCTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTTGAC TACTGGGGCCAAAGCACTACGGTCACCGTCTCCTCA
96	CD33 VL of F2	artificial	aa	DIVMTQSPDLSVSLGERTTINCKSSQSVLDSSTNKNLSLAWYQQKPGQPPKILLISWASTRESGIPD RFSGSGGTDFLTITIDSLQPEDSATYCYQQSAHFPIITFGQGRLEIK
97	CD33 LCDR1 of F2	artificial	aa	KSSQSVLDSSTNKNLSLA
98	CD33 LCDR2 of F2	artificial	aa	WASTRES
99	CD33 LCDR3 of F2	artificial	aa	QQSAHFPIIT
100	CD33 VL of F2	artificial	nt	GACATCGTGATGACACAGTCTCCAGACTCCCTGTCTGTGTCTCTGGGCGAGAGGACCACTCAAC TGCAAGTCCAGCCAGAGTGTTTTAGACAGCTCCACGAAATAAGAACTCCTTAGCTTGGTACCAGCAG AAACAGGACAGCTCCTAAATTAATCTCTTCTGGGCACTACGCGGGAATCCGGGATCCCTGAC CGATTCACTGGCAGCGGTCTGGGACAGATTTCATCTCACTATTGACAGCCTGCAGCCTGAAGAT TCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCAGATCACTTTGGCCAAAGGACACGACTG GAGATTAAA
101	CD33 HL of F2	artificial	aa	QVQLVQSGAEVKKPGEISVKVSKCASGYTFITNYGMNWVKQAPGQGLEWMGWINTYTGETNYADKFGQ RVFTTSDTSTSTAYMELRNILKSDDTAVYICARWSWSDGYVYFDYWGQGTITVTVSSGGGSGGGGS GGGSDIVMTQSPDLSVSLGERTTINCKSSQSVLDSSTNKNLSLAWYQQKPGQPPKILLISWASTRE SGIPDRFSGSGGTDFLTITIDSLQPEDSATYCYQQSAHFPIITFGQGRLEIK
102	CD33 HL of F2	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAACACCTACACTGGAGAGACAAAATATGCTGATAAGTTCCAGGGA CGGTTACCTTCACCTTCGATACCTTACCAGCACTGCCATAATGGAACCTCCGCAACCTCAAAAGT GATGACACGGCTGTATATTAATGCTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGCACTACGGTCACCGTCTCCTCAGGTGGTGGTTCCTGGCGGGCGGCTCC GGTGGTGGTGGTTCAGACATCGTGATGACACAGTCTCCAGACTCCCTGTCTGTGTCTCTGGCGAG AGGACCACTCAACTGCAAGTCCAGCCAGAGTGTTTTAGACAGCTCCACGAATAAGAACTCCTTA GCTTGGTACCAGCAGAAACCAGGACAGCTCCTAAATTAATCTCTTCCCTGGGCACTTACGCTATTGACAGC TCCGGGATCCCTGACCGGATTCAGTGGCAGCGGTCTGGGACAGATTTCATCTCACTATTGACAGC CTGACAGCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCAGTCACTCCCGATCACCTTTGGC CAAGGGACACGACTGGAGATTAAA

103	CD33 F2 HL x H2C HL	artificial	aa	QVQLVQSGAEVKKPESVKVSCKASGYTFITNYGMNVWKQAPGQGLEWMGWINTYTGETINYADKFKQG RVFTTSDTSTSTAYMELRNILKSDDTAVYYCARWSWSDGYVYFDYWGQGTTVTVSSGGGGSGGGGS GGGSDIVMTQSPDLSVSLGERTTINCKSSQSLDSSSTNKNLSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYYCQSAHFPITFFGQGRLEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAAASGFTFNKYAMNWRQAPGKGLEWVARIRSKYNNIYATYYADSVKGRFTISR DDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYISYWAYWGQGTILVTVSSGGGGSGGGSGGGGS QTVVTQEPISLTIVSPGGTVTLITCGSSSTGAVTSGYKPNWVQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGGKAALITLSGVQPEDEAEYYCALWYSNRWVFGGGTKLITVL
104	CD33 F2 HL x H2C HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTTGGAGAGTCAGTCAAGGTCTCCTGTC AAGGTAGCGGGTATACCTTCACAAAACATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACCTACACTGGAGAGACAAAATATGCTGATAAGTTCCAGGA CGGTTACCTTCACCTTCGGATACCTCTACCAGCACTGCCATATATGGAACCTCCGCAACCTCAAAAGT GATGACACGGCTGTATATTTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCACACGGTACCGTACCGTCTCCTCAGGTGGTGGTCTTGGCGGGCGGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGTCTGTGTCTCTGGCGGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCACGAATAAGAACTCCCTTA GCTTGGTACCAGAGAAAACAGGACAGCCTCCTAAATTTACTCCTTTCTCTGGGCATCTACGCGGGAA TCCGGGATCCCTGACCGATTCAGTGGCAGCGGTCTGGGACAGATTTCACTCTCACTATTGACAGC CTGCAGCCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCGATCACTCTTGGC CAAGGGACACAGCTGGAGATTAAATCCGGAGGTGGTGGCTCCGAGGTGACGTGGTCGAGTCTGGA GGAGGATTGGTGCAGCTGGAGGGTCAATTGAAAACCTCTCATGTGCAGCTCTTGATTCACCTTCAAT AAGTACGCCATGAACTGGGTCCGCCAGGCTCCAGGAAAAGGTTTGGAAATGGGTGTCTCGCATAGA AGTAAATATAATAATATGCAACATATTTATGCCGATTAGTGAAGACAGGTTACCATCTCCAGA GATGATTCAAAAACACCTGCCATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACCTTCGGTAAATAGCTACATATCTTACTTGGCTTACTTGGGGCCAAAGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGTCTTGGCGGCGCGGCTCCGGTGGTGGTGTCTT CAGACTGTTGTGACTCAGGAACCTTCACCTACCGTATCACCTGGTGGAACAGTCACACTCACTGT GGCTCCTCGACTGGGGCTGTTACATCTGGCTACTACCCAAACTGGGTCCAAACAAAAACAGGTCAG GCACCCCGTGGTCTAATAGGTGGGACTAAAGTTCTCGCCCCGGTACTCCTTGCAGATTCTCAGGC TCCCCTGCTTGGAGGCAAGGCTGCCCTTCACCCCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGCTCTATGGTACAGCAACCGCTGGGTGTTTCGGTGGAGGAACCAAACTGACTGTCTCTA
105	CD33 F2 HL x F12Q HL	artificial	aa	QVQLVQSGAEVKKPESVKVSCKASGYTFITNYGMNVWKQAPGQGLEWMGWINTYTGETINYADKFKQG RVFTTSDTSTSTAYMELRNILKSDDTAVYYCARWSWSDGYVYFDYWGQGTTVTVSSGGGGSGGGGS GGGSDIVMTQSPDLSVSLGERTTINCKSSQSLDSSSTNKNLSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSLQPEDSATYYCQSAHFPITFFGQGRLEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAAASGFTFNKYAMNWRQAPGKGLEWVARIRSKYNNIYATYYADSVKGRFTISR DDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYISYWAYWGQGTILVTVSSGGGGSGGGSGGGGS QTVVTQEPISLTIVSPGGTVTLITCGSSSTGAVTSGYKPNWVQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGGKAALITLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLITVL

106	CD33 F2 HL x F12Q HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACCTACACTGGAGAGACAAACTATGCTGATAAGTTCCAGGGA CGCGTTACCTTCACCTTCGGATACCTCTACCAGCACTGCCTATATGGAACCTCCGCAACCTCAAAAGT GATGACACGGCTGTATATTAATGCTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTTGAC TACTGGGGCCAAAGGCACTACGGTACCGTCTCCTCAGGTGGTGGTTCTGGCGGCGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGTCTGTGTCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCACGAATAAGAACTCCCTTA GCTTGGTACAGACAGAAAACAGGACAGCCTCCTAAAATTACTCTCTTCTGGGCATCTACGCGGGA TCCGGGATCCCTGACCGATTTCAGTGGCAGCGGTCTGGGACAGATTCTACTCTCACTATTGACAGC CTGACGCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCCACTTCCCGATCACTTTGGC CAAGGACACGACTGGAGATTAAATCCGAGGTGGTGGCTCCGAGGTGCAGTGGTCGAGTCTGGA GGAGGATTGGTCAGCTGGAGGTCATTGAAACTCTCATGTGCAGCTCTGGATTCACTTCAAT AGCTACGCCATGAACTGGGTCCGCCAGGCTCCAGGAAAAGGTTTGGAAATGGGTTCGCATTAAGA AGTAAATATAATAATTATGCAACATATTATGCCGATTTCAGTGAAGGAGGTTTCCCATCTCCAGA GATGATTCAAAAAACACTGCCTATCTACAAATGAACAACTTGAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAATAGCTACGTTTCTGGTGGCTTACTGGGGCCAAAGG ACTCTGGTCAACCGTCTCCTCAGGTGGTGGTGTCTGGCGGCGGCTCCGGTGGTGGTGTCT CAGACTGTGTGACTCAGGAACTTCACCTACCGTATCACCTGGTGGAAACAGTCACACTCACTTGT GGCTCTCGACTGGGCTGTACATCTGGCAACTACCCAACTGGGTCCAAACAAACACAGGTCAAG GCACCCGTGGTCTAAATAGGTGGACTAAGTTCTCGCCCCGCTACTCTTCCAGATTCTCAGGC TCCCTGCTTGGAGGCAAGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGTCTTATGTTACAGCAACCGTGGGTGTCTCGTGGAGAAACCAACTGACTGTCTCTA
107	CD33 F2 HL x I2C HL	artificial	aa	QVQLVQSGAEVKKPESVKVSKASGYTFTNYGMWVWYKQAPGQGLEWMGWINTYTGENTYADKFKQ RVFTTSDTSTSTAYMELRNILKSDDTAVYYCARWSWSDGYVYFDYWGQGTFTVTVSSGGGSGGGGS GGGSDIVMTQSPDSLVSISLGERTTINCKSSQSVLDSSTNKNLSLAWYQKPGQPPKLLISWASTRE SGIPDRFSGSGSDTFTLTIDSLQPEDSATYCYQQAHPITFTGQTRLEIKSGGGSEVQLVESG GGIVQPGGSLKLSCAASGFTFNKYAMNWRQAPGKLEWVARIRSKYNNYATYADSVKDRFTISR DDSKNTAYLQMNILKTEDTAVYYCVRHGNFGNSYISWAYWGQGTFTVTVSSGGGSGGGSGGGGS QTVVTOEPLSLTVSPGGFTVTLTCGSSTGAVTSGNYPNWYQKPGQAPRGLIGGTFKLAPGTPARFSG SLLGGKAALTLSGVQPEDEAEYYCVLWYSNRWVFGGTTKLTIVL
108	CD33 F2 HL x I2C HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACCTACACTGGAGAGACAAACTATGCTGATAAGTTCCAGGGA CGCGTTACCTTCACCTTCGGATACCTCTACCAGCACTGCCTATATGGAACCTCCGCAACCTCAAAAGT GATGACACGGCTGTATATTAATGCTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTTGAC TACTGGGGCCAAAGGCACTACGGTACCGTCTCCTCAGGTGGTGGTTCTGGCGGCGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGTCTGTGTCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCACGAATAAGAACTCCCTTA

				GCTTGGTACCAGCAGAAAAACAGGACAGCCTCCTAAATTACTCCTTTCTGGGCATCTACGGGGAA TCCGGGATCCCCTGACCGATTTCAGTGGCAGCGGTCTGGGACAGATTTCATCTCTACTATTGACAGC CTGCAGCCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCGATCACCCTTTGGC CAAGGACACAGACTGGAGATTAAATCCGAGGTGGTGGCTCCGAGGTGCAGTGGTCGAGTCTGGA GGAGGATTGGTGCAGCCTGGAGGGTCATTGAAACTCTCATGTGCAGCCTCTGGATTCACTTCAAT AAGTACGCCATGAACTGGGTCCGCGAGGCTCCAGGAAAGGGTTTGGAAATGGGTGCTCGCATAGA AGTAAATATAATAATTATGCAACATATATGCCGATTTCAGTGAAGACAGGTTACCATCTCCAGA GATGATTCAAAAAACACTGCCCTATCTACAAATGAACAACCTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAAATAGCTACATATCTCTACTGGGTTCCTGGGGCCAAGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGTCTTGGCGGCGGGCTCCGGTGGTGGTGTCT CAGACTGTTGTGACTCAGGAACTTCACTCACCGTATCACCTGGTGGAAACAGTCACACTCACTTGT GGCTCCTCGACTGGGGCTGTTACATCTGGCAACTACCCAAACTGGGTCCAAACAAAAACAGGTCAG GCACCCCGTGGTCTAAATAGGTGGGACTAAAGTTCTTCCGCCCCGGTACTCCTGCCAGATTCTCAGG TCCCCTGCTTGGAGGCAAGGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGTTCTATGGTACAGCAACCGCTGGGTGTTTCGGTGGAGGAACCAAACTGACTGTCTTA QVQLVQSGAEVKKPESVKVSKKASGYTFITNYGMNWVKQAPGQGLEWMGWINTYTGEPTVADKFFQG RVMTTDTSTSTAYMEIRNLRSDDTAVYYCARWSWSDDGYVYFDYWGQGTTVTVSS
109	CD33 VH of B10	artificial	aa	NYGMN
110	CD33 HCDR1 of B10	Artificial	aa	
111	CD33HCDR2 of B10	artificial	aa	WINTYTGEPTVADKFFQG
112	CD33 HCDR3 of B10	artificial	aa	WSWSDGYVYFDY
113	CD33 VH of B10	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGTGAGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGGTGGATAAACACCTACACTGGAGAGCCAACTATGCTGATAAGTTCAGGGA CGCTTACCATGACTACGGATACCTCTACCAGCACTGCCTATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCACACTACGGTCACCGTCTCCTCA
114	CD33 VL of B10	artificial	aa	DIVMTQSPDLSLTIVSLGERTTINCKSSQSVLDSNNKNSLAWYQQKPGQPPKLLLSWASTRESGIPD RFGSGSGTDFTLTIDGLQPEDSATYFCQQSAHFPIITFGQGTRLEIK
115	CD33 LCDR1 of B10	artificial	aa	KSSQSVLDSNNKNSLA
116	CD33 LCDR2 of B10	artificial	aa	WASTRES
117	CD33 LCDR3 of B10	artificial	aa	QQSAHFPIIT
118	CD33 VL of B10	artificial	nt	GACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTTGGGCGAGAGGACCACCATCAAC

119	CD33 HL of B10	artificial	aa	TGCAAGTCCAGCCAGAGTGTGTTTATAGACAGCTCCAACAATAAGAACTCCTTAGCTTGGTACCAGCAG AAACCAGGACAGCCTCCTAAATTAATCTCTTCTGGGCATCTACGCGGGAATCCGGGATCCCTGAC CGATTAGTGGCAGCGGTTCTGGGACAGATTTCATCTCTCACTATTGACGGCTGCAGCCTGAAGAT TCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCAGTACCTTTGGCCAAAGGACACAGACTG GAGATTAAA QVQLVQSGAEVKKPGEISVKVSKKASGYTFITNYGMNWKQAPGQGLEWMGWINTYTGEPTYADKFFQG RVMTTDTSTSTAYMEIRNLRSDDTAVYYCARWSWSDGYVYFDYWGQGTFTVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERFTINCKSSQSLDSSNNKNSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGGTDFLTITIDGLQPEDSATYFCQSSAHFPIITFGQGTREIK CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGTGAGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAACTATGGAATGAACCTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACTACACTGAGAGAGCAACCTATGCTGATAAGTTCCAGGGA CGCTTACCATGACTACGGATACCTCTACACAGCATGCCATATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGCGCAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTTCTTGGCGGCGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCAACAATAAGAACTCCTTA GCTTGGTACCAGAGAAACCAGGACAGCCTCCTAAATTAATCTCTTCTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTGAGTGGCAGCGGTTCTGGGACAGATTTCATCTCACTATTGACGGC CTGCAGCCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCAGTCACTCTTGGC CAAGGACACGACTGGAGATTAAA
120	CD33 HL of B10	artificial	nt	QVQLVQSGAEVKKPGEISVKVSKKASGYTFITNYGMNWKQAPGQGLEWMGWINTYTGEPTYADKFFQG RVMTTDTSTSTAYMEIRNLRSDDTAVYYCARWSWSDGYVYFDYWGQGTFTVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERFTINCKSSQSLDSSNNKNSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGGTDFLTITIDGLQPEDSATYFCQSSAHFPIITFGQGTREIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNWKQAPGKGLEWVARIRSKYNNYATYYADSVKDRFTISR DDSKNTAYLQMNNLKTEDTAVYYCVRHGNFNSYISYWAYWGQGTFTVTVSSGGGSGGGGSGGGGS QTVVTQEPSTLTVSPGGTFTLTTCGSTGAVTSGYYPNWKQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGGKAAALTLISGVQPEDEAEYFCALWYSNRWVFGGKLTIVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGTGAGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAACTATGGAATGAACCTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACTACACTGAGAGCAACCTATGCTGATAAGTTCCAGGGA CGCGTTACCATGACTACGGATACCTCTACACAGCATGCCATATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGCGCAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTTCTTGGCGGCGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCAACAATAAGAACTCCTTA GCTTGGTACCAGAGAAACCAGGACAGCCTCCTAAATTAATCTCTTCTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTGAGTGGCAGCGGTTCTGGGACAGATTTCATCTCACTATTGACGGC
121	CD33 B10 HL x H2C HL	artificial	aa	QVQLVQSGAEVKKPGEISVKVSKKASGYTFITNYGMNWKQAPGQGLEWMGWINTYTGEPTYADKFFQG RVMTTDTSTSTAYMEIRNLRSDDTAVYYCARWSWSDGYVYFDYWGQGTFTVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERFTINCKSSQSLDSSNNKNSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGGTDFLTITIDGLQPEDSATYFCQSSAHFPIITFGQGTREIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNWKQAPGKGLEWVARIRSKYNNYATYYADSVKDRFTISR DDSKNTAYLQMNNLKTEDTAVYYCVRHGNFNSYISYWAYWGQGTFTVTVSSGGGSGGGGSGGGGS QTVVTQEPSTLTVSPGGTFTLTTCGSTGAVTSGYYPNWKQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGGKAAALTLISGVQPEDEAEYFCALWYSNRWVFGGKLTIVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGTGAGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAACTATGGAATGAACCTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACTACACTGAGAGCAACCTATGCTGATAAGTTCCAGGGA CGCGTTACCATGACTACGGATACCTCTACACAGCATGCCATATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGCGCAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTTCTTGGCGGCGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCAACAATAAGAACTCCTTA GCTTGGTACCAGAGAAACCAGGACAGCCTCCTAAATTAATCTCTTCTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTGAGTGGCAGCGGTTCTGGGACAGATTTCATCTCACTATTGACGGC
122	CD33 B10 HL x H2C HL	artificial	nt	QVQLVQSGAEVKKPGEISVKVSKKASGYTFITNYGMNWKQAPGQGLEWMGWINTYTGEPTYADKFFQG RVMTTDTSTSTAYMEIRNLRSDDTAVYYCARWSWSDGYVYFDYWGQGTFTVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERFTINCKSSQSLDSSNNKNSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGGTDFLTITIDGLQPEDSATYFCQSSAHFPIITFGQGTREIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNWKQAPGKGLEWVARIRSKYNNYATYYADSVKDRFTISR DDSKNTAYLQMNNLKTEDTAVYYCVRHGNFNSYISYWAYWGQGTFTVTVSSGGGSGGGGSGGGGS QTVVTQEPSTLTVSPGGTFTLTTCGSTGAVTSGYYPNWKQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGGKAAALTLISGVQPEDEAEYFCALWYSNRWVFGGKLTIVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGTGAGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAACTATGGAATGAACCTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACTACACTGAGAGCAACCTATGCTGATAAGTTCCAGGGA CGCGTTACCATGACTACGGATACCTCTACACAGCATGCCATATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGCGCAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTTCTTGGCGGCGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGAG AGGACCAACATCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGCTCCAACAATAAGAACTCCTTA GCTTGGTACCAGAGAAACCAGGACAGCCTCCTAAATTAATCTCTTCTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTGAGTGGCAGCGGTTCTGGGACAGATTTCATCTCACTATTGACGGC

123	CD33 B10 HL x F12Q HL	artificial	aa	CTGCAGCCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACATTCCCAGATCACTTTGGC CAAGGGACACGACTGGAGATTAAATCCGAGGTGGTGGCTCCGAGGTGCAGTGGTTCGAGTCTGGA GGAGGATTGGTGCAGCTGGAGGGTCATTGAAACTCTCATGTGCAGCCTCTTGATTCACCTTCAAT AAGTACGCCATGAACCTGGGTCCGCCAGGCTCCAGGAAAGGTTTGGAAATGGTTCGATTCGATTAAGA AGTAAATATAATAATTATGCAACATATTATGCCGATTAGTGAAGACAGGTTACCATCTCCAGA GATGATTCAAAAAACACATGCCTATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGAACTTCGGTAATAGCTACATATCTTACTGGGCTTACTGGGGCAAGGG ACTCTGGTCAACCGTCTCCCTCAGGTGGTGGTGTCTTGGCGCGCGCGCTCCGGTGGTGGTTCCT CAGACTGTTGTGACTCAGGAACCTTCACTACCGTATCACTGGTGGAAACAGTCACTCACTTGT GGTCTCTCGACTGGGGCTGTACATCTGGCTACTACCAAACTGGGTCCAAACAAACACAGGTGAG GCACCCCGTGGTCTAAATAGGTGGACTAAGTCTCTGCCCCGGTACTCTGCCAGATTCTCAGGC TCCCTGCTTGGAGGCAAGGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGCTCTATGGTACAGCAACCGCTGGGTGTTCGGTGGAGGAACCAAACTGACTGTCCCTA QVQLVQSGAEVKKPESVKVSCKASGYTFNYGMNWVKQAPGQGLEWMGWINTYTGEPYADKFKQ RVMTTDTSTSTAYMEIRNLRSDDTAVYICARWSWDGYVYFDYWGQGTITVTVSSGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSVLDSNNKNSLAWYQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDGLQPEDSATYCYQQSAHFPITFGQGTRELEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNSYAMNWVRQAPGKGLEWVARIRSKNNYATYADSVKGRFTISR DDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYVSWWAYWGQGTITVTVSSGGGSGGGGSGGGGS QTVVTQEPSTLVSPGGITVTLTLCGSSSTGAVTSGNYPNWVQKPGQAPRGLIGGTFKFLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEYYCVLWYSNRWVFGGTGKLTIVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCTGGTGAGTCAGTCAGTCAAGGTCTCTCTGC AAGGTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGGTGGATAAAACACCTACACTGGAGAGCCAACTATGCTGATAAGTTCAGGGA CGGTTACCATGACTACGGATACCTCTACCAGACTGCCTATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTGTCTGCGCGCGCGGTCC GGTGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCAACCATCAACTGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAACAATAAGAACTCCCTTA GCTTGGTACAGAGAAAACAGGACAGCTCCCTAAATTACTCCTTCTCTGGCATCTACGCGGAA TCCGGGATCCCTGACCCGATTCACTGGCAGCGGTCTTGGGACAGATTCACTCTCACTATTGACGGC CTGCAGCCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCAGATCACTTTGGC CAAGGACACGACTGGAGATTAAATCCGGAGGTGGTGGTCCGAGGTGACGTTGGTTCGAGTCTGGA GGAGGATTGGTGCAGCTGGAGGGTCAATTGAAACTCTCATGTGCAGCTCTGGATTCACTTCAAT AGCTACGCCCATGAACTGGGTCCGCCAGGCTCCAGGAAAAGGTTTGGAAATGGGTTCGCTCGATAAGA AGTAAATATAATAATTATGCAACATATTATGCCGATTAGTGAAGAGCAGGTTACCATCTCCAGA GATGATTCAAAAAACACATGCCTATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACCTTCGGTAATAGCTACGTTTCTCTGGTGGGCTTACTGGGGCCAAAGG ACTCTGGTCAACCGTCTCCTCAGGTGGTGGTGTCTTGGCGCGCGCGGTCCGGTGGTGGTGGTTCCT
124	CD33 B10 HL x F12Q HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCTGGTGAGTCAGTCAGTCAAGGTCTCTCTGC AAGGTAGCGGGTATACCTTCACAAAATATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGGTGGATAAAACACCTACACTGGAGAGCCAACTATGCTGATAAGTTCAGGGA CGGTTACCATGACTACGGATACCTCTACCAGACTGCCTATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTGTCTGCGCGCGCGGTCC GGTGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCAACCATCAACTGCAAGTCCAGCCAGAGTGTCTTAGACAGCTCCAAACAATAAGAACTCCCTTA GCTTGGTACAGAGAAAACAGGACAGCTCCCTAAATTACTCCTTCTCTGGCATCTACGCGGAA TCCGGGATCCCTGACCCGATTCACTGGCAGCGGTCTTGGGACAGATTCACTCTCACTATTGACGGC CTGCAGCCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCAGATCACTTTGGC CAAGGACACGACTGGAGATTAAATCCGGAGGTGGTGGTCCGAGGTGACGTTGGTTCGAGTCTGGA GGAGGATTGGTGCAGCTGGAGGGTCAATTGAAACTCTCATGTGCAGCTCTGGATTCACTTCAAT AGCTACGCCCATGAACTGGGTCCGCCAGGCTCCAGGAAAAGGTTTGGAAATGGGTTCGCTCGATAAGA AGTAAATATAATAATTATGCAACATATTATGCCGATTAGTGAAGAGCAGGTTACCATCTCCAGA GATGATTCAAAAAACACATGCCTATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACCTTCGGTAATAGCTACGTTTCTCTGGTGGGCTTACTGGGGCCAAAGG ACTCTGGTCAACCGTCTCCTCAGGTGGTGGTGTCTTGGCGCGCGCGGTCCGGTGGTGGTGGTTCCT

125	CD33 B10 HL x I2C HL	artificial	aa	CAGACTGTTGTGACTCAGGAAACCTTCACTCACCGTATCACTGGTGGAAACAGTCACATCACTTGT GGCTCCTCGACTGGGGCTGTTACATCTGGCAACTACCCAAACTGGGTCCAAACAAAAACAGGTCAG GCACCCCGTGGTCTAAATAGGTGGGACTAAGTTCCTCGCCCCGGTACTCCTGCCAGATTCTCAGGC TCCCCTGCTTGGAGGCAAGGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGTTCTATGGTACAGCAACCGCTGGGTGTTTCGGTGGAGGAACCAAACTGACTGTCTCTA QVQLVQSGAEVKKPQESVKVSKASGYTFITNYGMNWKQAPGQGLEWMGWINTYTGEPTIADKFFQG RVITMTTDTSTSTAYMEIRNLRSDDTAVYYCARWSWSDGYVYFDYWGQGTITVTVSSGGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTITINCKSSQSLDSSNNKNSLAWYQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDGLQPEDSATYFCQOQSAHFPITFTGQGRLEIKSGGGGSEVQLVIESG GGILVQPGGSLKLSCAASGFTFNKYAMNWNVRQAPGKGLEWVARIRSKYNNYATYYADSVKDRFTISR DDSKNTAYLQMNLIKTEDTAVYYCVRHNFGNSYISYWAYWGQGLVTVSSGGGGSGGGSGGGGS QTVVTVQEPSTLVSPGGTVTLTTCGSSTGAVTSGNYPNWVQKPGQAPRGLIGGTFKFLAPGTPARFSG SLIIGKKAALTLSGVQPEDEAEYYCVLMYSNRWVFGGGTKLTVL
126	CD33 B10 HL x I2C HL	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAAGAGCCTGGTGAGTCAGTCAGTCAAGGTCTCTCTGC AAGGCTAGCGGGGTATACCTTTCACAAACTATGGAAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCAACCTATGCTGATAAAGTTCAGGGGA CGCGTTACCATGACTACGGATACCTCTACCAGCACTGCCATATATGGAATCCGCAACCTCAGAAAT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCATACGGTCACCGTCTCCTCAGGTGGTGGTGGTCTTGGCGGCGGGCTCC GGTGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCCCTGACTGTGTCTCTGGGCGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTTTTAGACAGTCCAAACAATAAGAACTCCTTTA GCTTGGTACCAGAGAAAACAGGACAGCTCCTCTAAATTTACTCTCTTCTCTGGGCACTACGCGGGAA TCCGGGATCCCCTGACCGATTTCAGTGGCAGCGGTTCTGGGACAGATTCACCTCTCACTATTGACGGC CTGCAGCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCCACTTCCCGATCACCTTTGGC CAAGGACACGACTGGAGATTAAATCCGGAGGTGGTGGCTCCGAGGTGCAGTGGTCGAGTGTGA GGAGGATTGGTGCAGCTGGAGGGTCAATTGAAACTCTCATGTGCAGCTCTGGATTCACTTCAAT AAGTACGCCATGAACCTGGGTCCGCCAGGCTCCAGGAAAGGTTTGGAAATGGGTGCTCGCATAGA AGTAAATATAATAATTATGCAACATATTATGCCGATTTCAGTGAAGACAGGTTCCCATCTCCAGA GATGATTCAAAAAACACTGCCATATCTACAAATGAACAACCTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAATAGCTACATATCTTACTGGGCTTACTGGGGCCAAGGG ACTCTGGTCAACCGTCTCCTCAGGTGGTGGTGGTCTTGGCGGCGCGGCTCCGGTGGTGGTGGTCT CAGACTGTTGTGACTCAGGAAACCTTCACTCACCGTATCACTGGTGGAAACAGTCACATCACTTGT GGCTCCTCGACTGGGGCTGTTACATCTGGCAACTACCCAAACTGGGTCCAAACAAAAACAGGTCAG GCACCCCGTGGTCTAAATAGGTGGGACTAAGTTCCTCGCCCCGGTACTCCTGCCAGATTCTCAGGC TCCCCTGCTTGGAGGCAAGGCTGCCCTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGTTCTATGGTACAGCAACCGCTGGGTGTTTCGGTGGAGGAACCAAACTGACTGTCTCTA
127	CD33 VH of E11	artificial	aa	QVQLVQSGAEVKKPQESVKVSKASGYTFITNYGMNWKQAPGQGLEWMGWINTYTGEPTIADKFFQG RVITMTTDTSTSTAYMEIRNLGDDDTAVYYCARWSWSDGYVYFDYWGQGTITVTVSS

128	CD33 HCDR1 of E11	artificial	aa	NYGMN	
129	CD33 HCDR2 of E11	artificial	aa	WINTYTGEPTYADKFGQ	
130	CD33 HCDR3 of E11	artificial	aa	WSWSDGYVYFDY	
131	CD33 VH of E11	artificial	nt	CAGGTGCAGCTGGTGCAGCTGGAGCTGAGGTGAAGAACCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAACACCTACACTGGAGAGCCAACTATGCTGATAAGTTCCAGGGA CGCGTTACCATGACTACGGATACCTCTACCAGCACTGCCTATATGGAATCCGCAACCTCGGAGGT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGCACTTCGGTCACCGTCTCCTCA	
132	CD33 VL of E11	artificial	aa	DIVMTQSPDLSLTIVSLGERTTINCKSSQSVLDSSTNKNLSLAWYQQKPGQPPPKLLLSWASTRESGIPD RFGSGSGTDFTLTIDSPQPEDSATYCYCQSAHFPIITFGQGRLEIK	
133	CD33 LCDR1 of E11	artificial	aa	KSSQSVLDSSTNKNLSLA	
134	CD33 LCDR2 of E11	artificial	aa	WASTRES	
135	CD33 LCDR3 of E11	artificial	aa	QQSAHFPIIT	
136	CD33 VL of E11	artificial	nt	GACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGGCGAGAGGACCAACATCAAC TGCAAGTCCAGCCAGAGTGTTTTAGACAGCTCCACGAAATAAGAACTCCTTAGCTTGGTACCAGCAG AAACAGGACAGCTCTCTAAATTACTCTCTTCTGGGCATCTACGCGGGAATCCGGGATCCCTGAC CGATTGAGTGGCAGCGGTCTGGGACAGATTTCACTCTCACTATTGACAGCCCGCAGCCTGAAGAT TCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCAGATCACTTTGGCCCAAGGACACGACTG GAGATTAAA	
137	CD33 HL of E11	artificial	aa	QVQLVQSGAEVKKPGEISVKVSKASGYTFITNYGMNWKQAPGQGLEWMGWINTYTGEPTYADKFGQ RVTMTTDTSTSTAYMEIRNLGGDDTAVVYCARWSWSDGYVYFDYWGQGTSTVTVSSGGGSGGGGS GGGSDIVMTQSPDLSLTIVSLGERTTINCKSSQSVLDSSTNKNLSLAWYQQKPGQPPPKLLLSWASTRE SGIPDRFSGSGGTDFLTITIDSPQPEDSATYCYCQSAHFPIITFGQGRLEIK	
138	CD33 HL of E11	artificial	nt	CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCTTGAGAGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAAACTATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAACACCTACACTGGAGAGCCAACTATGCTGATAAGTTCCAGGGA CGCGTTACCATGACTACGGATACCTCTACCAGCACTGCCTATATGGAATCCGCAACCTCGGAGGT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGCACTTCGGTCACCGTCTCCTCAGGTGGTGGTTCTTGGCGGCGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTTTATAGACAGCTCCACGAATAAGAACTCCTTA	

139	CD33 E11 HL x H2C HL	artificial	aa	GCTTGGTACCAGCAGAAAAACAGGACAGCCTCCTAAATTACTCCTTTCTTGGGCATCTACGCGGGAA TCCGGGATCCCTGACCGATTCACTGGCAGCGGTCTGGGACAGATTCTACTCTCACTATTGACAGC CCGACGCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCGATCACTTTGGC CAAGGACACGACTGGAGATTAAA QVQLVQSGAEVKKPGEISVKVSKASGYTFTNYGMNWKQAPGQGLEWMGWINTYTGEPTIADKFKQG RVITMTTDTSTSTAYMEIRNLGGDDTAVYICARWSWSDGYVYFDYWGQGTSTVTVSSGGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSVLSDSTNKNLSLAWYQKPKGPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSPQPEDSATYICQSAHFPITFGQGRLEIKSGGGSEVQLVESG GGLVQPGGSLKLSCAASGFTFNKYAMNWRQAPGKGLIEWVARIRSKYNNYATYYADSVKDRFTISR DDSKNTAYLQMNLIKTEDTAVYYCVRHGNFNSYISYWAYWGQGTLLVTVSSGGGGSGGGSGGGGS QTVVTEPSLTVSPGGTVTLITCGSSTGAVTSGYYPNWVQKPKGAPRGLIGGTFKLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
140	CD33 E11 HL x H2C HL	artificial	nt	CAGGTGACAGTGGTGCAGTCTGGAGCTGAGGTGAAGAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAAACTATGGAAATGAACCTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAACACCTACACTGGAGAGCAACCTATGCTGATAAGTTCCAGGGA CGCGTTACCATGACTACGGATACCTCTACCAGCACTGCCATATATGGAATCCGCAACCTCGAGGT GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGCACTTCGGTCACCGTCTCCTCAGGTGGTGGTGTCTGGCGCGCGGCTCC GGTGGTGGTGGTCTTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCAACCATCAACTGCAAGTCCAGCCAGAGTGTTTAGACAGTCCACGAATAAGAACTCCTTA GCTTGGTACAGCAGAAAAACAGGACAGCCTCCTAAATTTACTCTCTTTCTTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTCACTGGCAGCGGTCTGGGACAGATTCTACTCTCACTATTGACAGC CCGACGCTGAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCCACTTCCCGATCACTTTGGC CAAGGACACGACTGGAGATTAAATCCGGAGGTGGTGGCTCCGAGGTGCAGTGGTCGAGTCTGGA GGAGGATTGGTCAGCTGGAGGGTCAATTGAAACTCTCATGTGCAGCCTCTGGATTCACTTCAAT AAGTACGCCATGAACCTGGGTCCGCCAGGCTCCAGGAAAGGGTTTGGAAATGGGTGTCTCGCATAGA AGTAAATATAATAATTATGCAACATATTATGCCGATTCACTGAAAGACAGGTTCCCATCTCCAGA GATGATTCAAAAAACACTGCCATATCTACAAATGAACAACTTGAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAATAGCTACATATCTTACTGGGCTTACTGGGGCCAAAGG ACTCTGGTCAACCGTCTCCTCAGGTGGTGGTGTCTTGGCGGCGGCTCCGGTGGTGGTGTCT CAGACTGTGTGACTCAGGAACTTCACTCACCGTATCACCTGGTGGAAACAGTCACTTGTGT GGCTCTCGACTGGGGCTGTACATCTGGCTACCTCAAACTGGGTCCAAACAAACAGGTCAG GCACCCGCTGGTCTAAATAGGTGGGACTAAGTTCTCTCCCCCGGTCTCTGCGAGATTCTCAGGC TCCCTGCTTGGAGGCAAGGCTGCCCTCACCTCTCAGGGTACAGCCAGAGGATGAGGACAGAAAT TACTGTGCTCTATGTTACAGCAACCGTGGGTGTTCGGTGGAGGAAACCAACTGACTGTCTTA
141	CD33 E11 HL x F12Q HL	artificial	aa	QVQLVQSGAEVKKPGEISVKVSKASGYTFTNYGMNWKQAPGQGLEWMGWINTYTGEPTIADKFKQG RVITMTTDTSTSTAYMEIRNLGGDDTAVYICARWSWSDGYVYFDYWGQGTSTVTVSSGGGGSGGGGS GGGSDIVMTQSPDSLTVSLGERTTINCKSSQSVLSDSTNKNLSLAWYQKPKGPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSPQPEDSATYICQSAHFPITFGQGRLEIKSGGGSEVQLVESG

142	CD33 E11 HL x F12Q HL	artificial	nt	GGLVQPGGSLKLSCAAASGFTFNSYAMNWWVQAPGKGLEWVARIRSKYNNIYATYYADSVKGRFTISR DDSKNTAYILQMNILKTEDTAVYYCVRHGNFNSYVSWWAYWQGTLVTVSSGGGSGGGSGGGGS QTVVTTQEPSTLVSPGGTVTLTTCGSSTGAVTSGNYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGKKAALTLISGVQPEDEAEFYCVLMYSNRWVFGGGTKLTVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAATGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCAACCTATGCTGATAAGTTCCAGGGA CGGTTACCATGACTACGGATACCTCTACAGCACTGCCTATATGAAAATCCGAAACCTCGGAGGT GATGACACGGCTGTATATTAATGTGCGCGCTGGAGTTGAGTGATGGTTACTACTGTTTACTTTGAC TACTGGGGCCAAAGCACTTCGGTCACCGTCTCTCAGGTGGTGGTGGTTCCTGGCGGCGGGCTCC GGTGGTGGTGGTTCGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGCTGCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTTTTAGACAGCTCCACGAAATAAGAACTCCTTA GCTTGGTACCAGAGAAAACAGGACAGCTCCTAAAATTACTCTCTTCTGGGCATCTACGCGGGAA TCCGGGATCCCTGACCGATTGAGTGGCAGCGGGTCTGGGACAGATTCTACTCTCACTATTGACAGC CCGAGCCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCCACTTCCCGATCACTTTGGC CAAGGGACACGACTGGAGATTAAATCCGAGGTGGTGGCTCCGAGGTGCAGCTTGAGTTCACCTTCAAT GGAGGATTGGTGCAGCTGGAGGGTCATTGAAAATCTCATGTGCAGCCTCTGGATTCACTTCAAT AGCTACGCCATGAACCTGGGTCCGCGCAGGCTCCAGGAAAGGGTTTGGAAATGGGTGCTCGCATAAAGA AGTAAATATAATAATATGCAACATATATATGCCGATTGATGAAAGCAGGTTACCATCTCCAGA GATGATTCAAAAAACACTGCCCTATCTACAAATGAACAACTTGAAAACTGAGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAAATAGCTACGTTTCTTGGTGGCTTACTTGGGCGCAAGGG ACTCTGGTCACCGTCTCTCAGGTGGTGGTGGTCTTGGCGGCGGGCTCCGGTGGTGGTGGTCTT CAGACTGTTGTGACTCAGGAACTTCACTCACCGTATCACTGGTGGAAACAGTCACACTCACTTGT GGCTCTCGACTGGGGCTGTTACATCTGGCAACTACCCAACTGGGTCCAAACAAAACCCAGGTCAG GCACCCCGTGGTCTAAATAGGTGGGACTAAAGTTCTCGCCCCGGTACTCTTCCAGATTCTCAGGC TCCCCTGCTTGGAGGCAAGGCTGCCCTCACCCCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGTTCTATGGTACAGCAACCGCTGGGTGTTCTGGTGGAGGAACCAAACTGACTGTCTCTA QVQLVQSGAEVKKPGEISVKVSKCASGYTFTNYGMNWKQAPGQGLEWMGWINTYTGEPTYADKFFQG RVTMTTDTSTSTAYMEIRNLGGDDTAVYYCARWSWSDGYVYFDYWQGQTSVTVSSGGGSGGGGS GGGSDDIVMTQSPDSLTVSLGERTTINCKSSQSVLDSSTNKNLSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSPQPEDSATYCYQQSAHFPIITFGQGRLEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAAASGFTFNKYAMNWWVQAPGKGLEWVARIRSKYNNIYATYYADSVKDRFTISR DDSKNTAYILQMNILKTEDTAVYYCVRHGNFNSYISYWAYWQGTLVTVSSGGGSGGGSGGGGS QTVVTTQEPSTLVSPGGTVTLTTCGSSTGAVTSGNYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGKKAALTLISGVQPEDEAEFYCVLMYSNRWVFGGGTKLTVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAATGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCAACCTATGCTGATAAGTTCCAGGGA CGCGTTACCATGACTACGGATACCTCTACAGCACTGCCTATATGAAAATCCGCAACCTCGGAGGT
143	CD33 E11 HL x I2C HL	artificial	aa	QVQLVQSGAEVKKPGEISVKVSKCASGYTFTNYGMNWKQAPGQGLEWMGWINTYTGEPTYADKFFQG RVTMTTDTSTSTAYMEIRNLGGDDTAVYYCARWSWSDGYVYFDYWQGQTSVTVSSGGGSGGGGS GGGSDDIVMTQSPDSLTVSLGERTTINCKSSQSVLDSSTNKNLSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSPQPEDSATYCYQQSAHFPIITFGQGRLEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAAASGFTFNKYAMNWWVQAPGKGLEWVARIRSKYNNIYATYYADSVKDRFTISR DDSKNTAYILQMNILKTEDTAVYYCVRHGNFNSYISYWAYWQGTLVTVSSGGGSGGGSGGGGS QTVVTTQEPSTLVSPGGTVTLTTCGSSTGAVTSGNYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGKKAALTLISGVQPEDEAEFYCVLMYSNRWVFGGGTKLTVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAATGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCAACCTATGCTGATAAGTTCCAGGGA CGCGTTACCATGACTACGGATACCTCTACAGCACTGCCTATATGAAAATCCGCAACCTCGGAGGT
144	CD33 E11 HL x I2C HL	artificial	nt	QVQLVQSGAEVKKPGEISVKVSKCASGYTFTNYGMNWKQAPGQGLEWMGWINTYTGEPTYADKFFQG RVTMTTDTSTSTAYMEIRNLGGDDTAVYYCARWSWSDGYVYFDYWQGQTSVTVSSGGGSGGGGS GGGSDDIVMTQSPDSLTVSLGERTTINCKSSQSVLDSSTNKNLSLAWYQQKPGQPPKLLLSWASTRE SGIPDRFSGSGSGTDFTLTIDSPQPEDSATYCYQQSAHFPIITFGQGRLEIKSGGGGSEVQLVESG GGLVQPGGSLKLSCAAASGFTFNKYAMNWWVQAPGKGLEWVARIRSKYNNIYATYYADSVKDRFTISR DDSKNTAYILQMNILKTEDTAVYYCVRHGNFNSYISYWAYWQGTLVTVSSGGGSGGGSGGGGS QTVVTTQEPSTLVSPGGTVTLTTCGSSTGAVTSGNYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGKKAALTLISGVQPEDEAEFYCVLMYSNRWVFGGGTKLTVL CAGGTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGC AAGGCTAGCGGGTATACCTTCACAAAATATGGAATGAATGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGCTGGATAAAACACCTACACTGGAGAGCAACCTATGCTGATAAGTTCCAGGGA CGCGTTACCATGACTACGGATACCTCTACAGCACTGCCTATATGAAAATCCGCAACCTCGGAGGT

				GATGACACGGCTGTATATTACTGTGCGCGCTGGAGTTGGAGTGAATGGTTACTACGTTTACTTTTGAC TACTGGGGCCAAAGGCACTTCGGTCAACCGTCTCCTCAGGTGGTGGTTCTTGGCGGCGGGCTCC GGTGGTGGTGGTTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTTTTAGACACAGCTCCACGAAATAAGAACTCCTTTA GCTTGGTACCAGACGAAAAACAGGACAGCTCTTAAATTACTCTCTTCTTGGGCACTACGCGGGAA TCCGGGATCCCTGACCGATTCAGTGGCAGCGGGTCTGGGACAGATTTCACTCTCACTATTGACAGC CCGAGCCTGAAAGATTCTGCAACTTACTATTGTCAACAGTCTGCCACTTCCCGATCACTTTTGGC CAAGGACACGACTGGAGATTAAATCCGGAGGTGGTGGCTCCGAGGTGCAGCTGGTCGAGTCTGGA GGAGGATTGGTGCAGCCTGGAGGGTCATTGAAAACTCTCATGTGCAGCCTCTTGGATTCACTTCAAT AAGTACGCCATGAACCTGGGTCCGCCAGGTCCAGGAAAGGGTTTGGAAATGGGTGCTCGCATTAAGA AGTAAATATAATAATTATGCAACATATTATGCCGATTAGTGAAAGACAGGTTTACCATCTCCAGA GATGATTCAAAAAACACTGCCATATCTACAAATGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAAATAGTACATATCTTACTTGGGCTTACTTGGGGCCAAAGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGTCTTGGCGGCGGGCTCCGGTGGTGGTGGTTCT CAGACTGTTGTGACTCAGGAACCTTCACTCAACCGTATCACTGGTGGAAACAGTCACTCACTTGT GGCTCCTCGACTGGGGCTGTTACATCTGGCAACTACCAAACTGGGTCCAAAAACCAAGTCAG GCACCCCGTGGTCTAAATAGGTGGGACTAAGTTCTCGCCCCGGTACTCTTGCAGATTCTCAGGC TCCCTGCTTGGAGGCAAGGCTGCCCTCACCCCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATAT TACTGTGTTCTATGGTACAGCAACCGCTGGGTGTTCGGTGGAGGAACCAAACTGACTGTCTCTA
145	CD33	human	nt	ATGGGATGGAGCTGTATCATCTCTTCTTGGTAGCAACAGCTACAGGTGTACACTCCGATCCAAAT TTCTGGCTGCAAGTGCAGGAGTCAGTGACGGTACAGGAGGGTTTGTGCGTCTCTGTGCCCCTGCACT TTCTTCCATCCCATACCTTACTACGACAAGAACTCCCCAGTTTCATGGTTACTTGGTTCCGGGAAGGA GCCATTATATCCGGGACTCTCCAGTGGCCACAAACAACTAGATCAAGAACTACAGGAGGAGACT CAGGCGAGATTCCGCCCTCTTGGGGATCCCAGTAGGAAACAACTGCTCCCTGAGCATCTGTAGACGCC AGGAGGAGGATAATGGTTTCACTTCTTTTCGGATGGAGAGAGGAAAGTACCAAAATACAGTTTACAAA TCTCCCCAGCTCTCTGTGCATGTGACAGACTTGACCCACAGGCCCAAAATCTCTCATCCCTGGCACT CTAGAACCCGGCCACTCCAAAAACCTGACCTGCTCTGTGTCTTGGGCTGTGAGCAGGGAACACACC CCGATCTTCTCCTGGTTGTGAGCTGCCCCACCTCCTTGGGCCCCAGGACTACTCACTCCTCGGTG CTCATAATCACCCACGGCCCCAGGACCACGGCACCAACTGACCTGTGAGGTGAAAGTTTCGTGGA GCTGGTGTGACTACGGAGAGAACTCCAGCTCAACGTCACTTATGTTCCACAGAAACCCCAACAACT GGTATCTTTCCAGGAGATGGCTCAGGGAACCAAGAGACCACAGCAGGAGTGGTTTCATGGGGCCATT GGAGGAGCTGGTGTACAGCCCTGCTGCTCTTTTGTCTTGCCTCATCTTCTTTCATAGTGAAGACC CACAGGAGGAAAAGCAGCCAGGACAGCAGTGGGAGGAAATGACACCCACCTTACCACAGGTCAGCC TCCCCGAAACACACAGAAAGTCCAAGTTTACATGGCCCCACTGAAAACTCAAGCTGTTTCAGGTGCC GCCCCCTACTGTGGAGATGGATGAGGAGCTGCATTATGCTTCCCTCAAACCTTTCATGGGATGAATCCT TCCAAGGACACCTCCACCGAATACTCAGAGGTGAGGACCCAGTCCGGGCATCATCACCATCATCAT TGA
146	CD33	human	aa	MGWSCIILFLVATATGVHSDPNFWLQVQESVTYQEGLCVLVPCTFFHPIPYDYDKNSPVHGYWVFREG AIIISGDSPVATNKLIDQEVQEETQGRFRLLGDP SRNNCSLSIVDARRRDNGSYFFRMERGSTKYSYK

[illegible]

150	CD33 UD F12Q HL x AF5 HL	artificial	aa	GGTACTCCTGCCAGATTCTCAGGCTCCCTGCTTGGAGGCAAGGCTGCCCTCACCCCTCTCAGGGGTA CAGCCAGAGGATGAGGCAGAAATATTACTGTGCTCTATGGTACAGCAACCGTGGGTGTTTCGGTGA GGAACCAAACTGACTGTCTTATCCGGAGGTGGTGGCTCCAGGTGCAGTGGTCCAGTCTGGAGCT GAGGTGAAGAAAGCCTGGAGCGTCAGTCAAGGTCTCCTGCAAGGCTAGCGGTATACCTTCACAAAC TATGGAATGAACCTGGGTGAAGCAGGCTCCAGGACAGGGTTTAAAGTGGATGGGCTGGATAAACACC TACACTGGAGAGCCAAACATATGCTGATGACTTCAAGGACGGGTACCATGACTTCGGATACCTCT ACCAGCACTGCCCTATTTGGAACCTCCACAACTCAGAACTGATGACACGGCTGTATATTACTGTGCG CGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGACTACTTGGGCGCAAGGCAC'TACGGTCAAC GTCTCCTCAGGTGGTGGTTCTTGGCGCGCGGCTCCGGTGGTGGTGGTCTTGACATCGTGATG ACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGGCGAGAGGACCACTCAACTGCAAGTCCAGC CAGAGTGTTTTAGACAGCTCCAAGAAATAAGAACTCCTTAGCTTGGTACACAGCAAAACCCAGGACAG CCTCCTAAATTTACTCCTTCTTGGCATCTACGCGGGAATCCGGGATCCCTGACCGATTCTAGTGGC AGCGGTCTGGGACAGATTTCACTCTCACTATTGACAGCTGCAGCTGAAGATTCTGCAACTTAC TATTGTCAACAGTCTGCCCACTTCCCAGATCACCTTTGGCCAAAGGACACGACTGGAGATTAAA EVQLVESGGGLVQPGGSLKLSCAAAGFTTFNSYAMNWVRQAPKGLEWVARIRSKYNNIYATYYADSV KGRFTISRDDSKNTAYLQMNNLKLTEDTAVYVCVRHGNFNGSYVSWWAYWQGTLIVTVSSGGGSGG GGSGGGGQTVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGNYPNWQQKPGQAPRGLIGGTFKFLAP GTPARFSGSLLGKKAALTLSGVQPEDEAEYCVLMYSNRWVFGGTKLTVLSGGGGSQVQLVQSGA EVKKPGASVKVSKASGYTFNRYGMNWVKQAPGQGLKWMGWINITYTEPTYADDFKGRVTMTSDTS TSTAYLELHNLRSDDTAVYYCARWSWSDGYVYFDYWQGTITVTVSS9999s9999sDIVM TQSPDSLTVSLGERTTINCKSSQSVLDSKNKNSLAWYQQKPGPPKLLLSWASTRESGIPDRFSG SGSGTDFTLTIDSLQPEDSATYYCQSAHFPITFGQGRLEIK
151	CD33 UD F12Q HL x AF5 HL	artificial	nt	GAGGTGCAGCTGGTCGAGTCTGGAGGAGGATTGGTGCAGCTTGAGGGTCATTGAAAAC'TCTCATGT GCAGCCTCTGGATTCACTTCAATAGCTACGCCATGAAC'TGGGTCCGCCAGGCTCCAGGAAAAGGT TTTGAATGGGTGCTCGCATAGAAAGTAAATATAATTAATATGCAACATATATGCCGATTCACTG AAAGGCAGGTTCAACATCTCCAGAGATGATTCAAAAAACACTGCC'TATCTACAAATGAACAACTTG AAAACTGAGGACACTGCCGTGTACTGTGTGAGACATGGGAACTTCGGTAATAGCTACGTTTCC TGGTGGGCTTACTGGGGCCAAAGGACTCTTGGTCACCGTCTCCTCAGGTGGTGGTCTTGCGGC GGCGGCTCCGGTGGTGGTTCTCAGACTGT'TGTGACTCAGGAACTTCACTCACCGTATCACCT GGTGAACAGTCACTCACTTGTGGTCTCTCGACTGGGGTGT'TACATCTGGCAACTACCCAAAC TGGGTCCAAACAAAAACAGGTCAAGCACCCCGTGGTCTAA'TAGGTGGGACTAAGTTCCTCGCCCC GGTACTCTTCCAGATTCTCAGGCTCCCTGT'TGGAGGCAAGGTCGCCCTCACCTCTCAGGGGTA CAGCCAGAGGATGAGGCAGAAATATTACTGTGT'TATGGTACAGCAACCGTGGGTGTTTCGGTGA GGAACCAAACTGACTGTCTTATCCGGAGGTGGTGGCTCCCGGTGCAGCTGGTCCAGTCTGGAGCT GAGGTGAAGAAAGCCTGGAGCGTCAAGGTCTCCTGCAAGGCTAGCGGTATACCTTCACAAAC TATGGAATGAACCTGGGTGAAGCAGGCTCCAGGACAGGGTTTAAAGTGGATGGGCTGGATAAACACC TACACTGGAGAGCCAAACATATGCTGATGACTTCAAGGACGGGTACCATGACTTCGGATACCTCT

				ACCAGCACTGCCATTATTTGGAACCTCCACAACTCAGAACTGATGACACGGCTGTATATTACTGTGCG CGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGACTACTTGGGGCCAAAGCACTACGGTCACC GTCCTCAGGTGGTGGTTCTTGGCGCGCGGCTCCGGTGGTGGTTCTTGACATCGTGATG ACACAGTCTCCAGACTCCCTGACTGTGTCTCTTGGCGGAGAGGACCACCATCAACTGCAAGTCCAGC CAGAGTGTTTTAGACAGCTCCAAGAAATAAGAACTCCTTAGCTTGGTACCAGAGAAACCCAGGACAG CCTCCTAAATTAATCTCTTCTTGGGCATCTACGCGGAATCCGGGATCCCTGACCGATTCTGCAACTTAC AGCGGGTCTGGGACAGATTTCACTCTCACTATTGACAGCTGCAGCTGAAGATTCTGCAACTTAC TATTGTCAACAGTCTGCCCACTTCCCCGATCACCTTTGGCCAAAGGACACGACTGGAGATTAAA
152	CD33 UD I2C HL x AF5 HL	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNKYAMNWRQAPKGLIEWARIRSKYNNYATYYADSV KDRFTISRDDSKNTAYLQMNLIKTEDTAVYYCVRHGNFGNSYISYWAYWGQGLVTVSSGGGSGG GGSGGGSQTVVTTQEPSTVSPGGTVTLTCSSTGAVTSNYPNWVQKPGQAPRGLIGGTFKFLAP GTPARFSGSLIGGKAALTLSGVQPEDEAEYYCVLWYSNRWVFGGTTKLTVLSGGGGSQVQLVQSGA EVKKPGASVKVSKKASGYTFNRYGMNWVKQAPGQGLKWMGWINTYTGEPTYADDFKGRVTMTSDTS TSTAYLELHNLRSDDTAVYYCARWSWSDGYVYFDYWQGGTIVTVSS9999s9999sDIVM TQSPDSLITVSLGERTTINCKSSQSVLDSKKNKNSLAWYQKPGQPPKLLLSWAISTRESGIPDRFSG SGSGTDFTLITIDSLQPEDSATYCYCQSAHFPIITFGQGTRLEIK
153	CD33 UD I2C HL x AF5 HL	artificial	nt	GAGGTGACAGCTGGTCGAGTCTGGAGGAGGATTTGGTGACGCTTGAGGGTCAATTGAAAATCTCATGT GCAGCCTCTGGATTCACTTCAATAAGTACGCCATGAATGGGTCCGCCAGGCTCCAGGAAAAGGT TTGGAATGGGTGGTTCGCATAAAGAGTAAATATAATAATATGCAACATATATGCGGATTCAGTG AAAGACAGGTTACCATCTCCAGAGATGATTCAAAAAACACTGCCATCTACAAAATGAACAACTTG AAAACTGAGGACACTGCCGTGTACTACTGTGTGAGACATGGGAACTTCGGTAAATAGCTACATATCC TACTGGGCTTACTGGGGCCAAAGGACTCTGGTCACCGTCTCCTCAGGTGGTGGTTCTTGGCGGC GGCGGCTCCGGTGGTGGTTCTCAGACTGTTGTGACTCAGGAACTTCACCTACCGTATCACCT GGTGAAACAGTCACACTCACTTGTGGCTCCTCGACTGGGGCTGTTACATCTGGCAACTACCCAAAC TGGGTCCAACAAAAACAGGTCAGGCACCCCGTGGTCTAATAGGTGGGACTAAGTCTCGCCCCC GGTACTCCTGCCAGATTCTCAGGCTCCCTGCTTGGAGGCAAGGCTGCCCTCACCTCTCAGGGGTA CAGCCAGAGGATGAGGCAGAATATTACTGTGTTCTATGGTACAGCAACCGCTGGGTGTTTCGGTGGA GGAACCAAACTGACTGTCTTATCCGGAGGTGGTGGTCCAGGTGCAGCTGGTCCAGTCTGGAGCT GAGGTGAAGAAAGCCTGGAGCGTCAGTCAAGGTCTCCTTGAAGGCTAGCGGGTATACCTTCACAAAC TATGGAATGAACCTGGGTGAAGCAGGCTCCAGGACAGGTTTAAAGTGGATGGGTGGATGAACACAC TACACTGGAGAGCCAAACATATGCTGATGACTTCAAGGACGGGTTACCATGACTTCGGATACCTCT ACCAGCACTGCCATTATTTGGAACCTCCACAACTCAGAACTGACACGGTGTATATATTACTGTGCG CGCTGGAGTTGGAGTGATGGTTACTACGTTTACTTTGACTACTTGGGGCCAAAGGCACTACGGTCACC GTCCTCCTCAGGTGGTGGTTCTTGGCGCGCGGCTCCGGTGGTGGTTCTTGACATCGTGATG ACACAGTCTCCAGACTCCCTGACTGTGTCTCTTGGGCGAGAGGACCACCATCAACTGCAAGTCCAGC CAGAGTGTTTTAGACAGCTCCAAGAAATAAGAACTCCTTAGCTTGGTACCAGAGAAACCCAGGACAG CCTCCTAAATTAATCTCTTCTTGGGCATCTACGCGGGAATCCGGGATCCCTGACCGATTCTCAGTGCG

				AGCGGGTCTGGGACAGATTTCACTCTCACTATTGACAGCCTGCAGCCTGAAGATTCTGCAACTTAC TATTGTCAACAGTCTGCCACTTCCCGATCCACTTTGGCCAAAGGACACGACTGGAGATTAAA
154	VL of F6A	artificial	aa	QTVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGGKAAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
155	VH of F6A	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAAASGFTFNIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KSRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNGFNSYVSFFAYWGQGTILVTVSS
156	VH-VL of F6A	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAAASGFTFNIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KSRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNGFNSYVSFFAYWGQGTILVTVSSGGGSGG GGGGGGSTVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAP GTPARFSGSLLGGKAAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
157	VL-P of F6A	artificial	aa	ELVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGGKAAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
158	VH-P of F6A	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAAASGFTFNIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KSRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNGFNSYVSFFAYWGQGTILVTVSS
159	VH-VL-P of F6A	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAAASGFTFNIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KSRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNGFNSYVSFFAYWGQGTILVTVSSGGGSGG GGGGGGSELVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAP GTPARFSGSLLGGKAAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
160	VL of H2C	artificial	aa	QTVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGGKAAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
161	VH of H2C	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAAASGFTFNKYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KDRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNGFNSYISYWAYWGQGTILVTVSS
162	VH-VL of H2C	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAAASGFTFNKYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KDRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNGFNSYISYWAYWGQGTILVTVSSGGGSGG GGGGGGSTVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAP GTPARFSGSLLGGKAAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
163	VL-P of H2C	artificial	aa	ELVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGGKAAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
164	VH-P of H2C	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAAASGFTFNKYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KDRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNGFNSYISYWAYWGQGTILVTVSS
165	VH-VL-P of H2C	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAAASGFTFNKYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KDRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNGFNSYISYWAYWGQGTILVTVSSGGGSGG GGGGGGSELVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAP GTPARFSGSLLGGKAAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
166	VL of H1E	artificial	aa	QTVVTQEPSTLVSPGGTVTLTCGSSTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGGKAAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
167	VH of H1E	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAAASGFTFNKYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KGRFTISRDDSKNTAYLQMNILKTEDTAVYYCVRHGNGFNSYISYWAYWGQGTILVTVSS

168	VH-VL of H1E	artificial	aa	EVQLVESGGGLEQPGGSLKLSCAASGFTFNISYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSV KGRFTISRDDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYLSFWAYWGQGTILVTVSSGGGSGG GGSGGGGQTVVTQEPSTLVSPGGTVTLTTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGGTKFLAP GTPARFSGSLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
169	VL-P of H1E	artificial	aa	ELVVTQEPSTLVSPGGTVTLTTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGGTKFLAPGTPARFSG SLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
170	VH-P of H1E	artificial	aa	EVQLVESGGGLEQPGGSLKLSCAASGFTFNISYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSV KGRFTISRDDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYLSFWAYWGQGTILVTVSS
171	VH-VL-P of H1E	artificial	aa	EVQLVESGGGLEQPGGSLKLSCAASGFTFNISYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSV KGRFTISRDDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYLSFWAYWGQGTILVTVSSGGGSGG GGSGGGSELVVTQEPSTLVSPGGTVTLTTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGGTKFLAP GTPARFSGSLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
172	VL of G4H	artificial	aa	QTVVTQEPSTLVSPGGTVTLTTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGGTKFLAPGTPARFSG SLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
173	VH of G4H	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNRYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSV KGRFTISRDDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYLSFWAYWGQGTILVTVSS
174	VH-VL of G4H	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNRYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSV KGRFTISRDDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYLSFWAYWGQGTILVTVSSGGGSGG GGSGGGGQTVVTQEPSTLVSPGGTVTLTTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGGTKFLAP GTPARFSGSLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
175	VL-P of G4H	artificial	aa	ELVVTQEPSTLVSPGGTVTLTTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGGTKFLAPGTPARFSG SLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
176	VH-P of G4H	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNRYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSV KGRFTISRDDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYLSFWAYWGQGTILVTVSS
177	VH-VL-P of G4H	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNRYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSV KGRFTISRDDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYLSFWAYWGQGTILVTVSSGGGSGG GGSGGGSELVVTQEPSTLVSPGGTVTLTTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGGTKFLAP GTPARFSGSLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
178	VL of A2J	artificial	aa	QTVVTQEPSTLVSPGGTVTLTTCRSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPSGTPARFSG SLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
179	VH of A2J	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNRYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSV KGRFTISRDDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYLSFWAYWGQGTILVTVSS
180	VH-VL of A2J	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNRYAMNWVRQAPGKGLEWVARIRSKYNNIATYYADSV KGRFTISRDDSKNTAYLQMNNLKTEDTAVYYCVRHGNFGNSYLSFWAYWGQGTILVTVSSGGGSGG GGSGGGGQTVVTQEPSTLVSPGGTVTLTTCRSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPS GTPARFSGSLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
181	VL-P of A2J	artificial	aa	ELVVTQEPSTLVSPGGTVTLTTCRSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPSGTPARFSG SLGGKAALTLTSGVQPEDEAEYICALWYSNRWVFGGGTKLTVL

182	VH-P of A2J	artificial	aa	EVQLLESGGGLVQPFGSLKLSCAAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KKRFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYLSWYAYWGQGTLLVTVSS
183	VH-VL-P of A2J	artificial	aa	EVQLLESGGGLVQPFGSLKLSCAAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KKRFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYLSWYAYWGQGTLLVTVSSGGGSGG GGSGGGSELVVTQEP SLTVSPGGT VTLTCRSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPS GTPARFSGSLGGKAAALTL SGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
184	VL of E1L	artificial	aa	QTVVTQEP SLTVSPGGT VTLTCRSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPSGTPARFSG SLGGKAAALTL SGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
185	VH of E1L	artificial	aa	EVQLVESGGGLVQPFGSLKLSCAAASGFTFNKIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KSRFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYTSYAYWGQGTLLVTVSS
186	VH-VL of E1L	artificial	aa	EVQLVESGGGLVQPFGSLKLSCAAASGFTFNKIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KSRFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYTSYAYWGQGTLLVTVSSGGGSGG GGSGGGQTVVTQEP SLTVSPGGT VTLTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPS GTPARFSGSLGGKAAALTL SGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
187	VL-P of E1L	artificial	aa	ELVVTQEP SLTVSPGGT VTLTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPSGTPARFSG SLGGKAAALTL SGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
188	VH-P of E1L	artificial	aa	EVQLLESGGGLVQPFGSLKLSCAAASGFTFNKIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KSRFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYTSYAYWGQGTLLVTVSS
189	VH-VL-P of E1L	artificial	aa	EVQLLESGGGLVQPFGSLKLSCAAASGFTFNKIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KSRFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYTSYAYWGQGTLLVTVSSGGGSGG GGSGGGSELVVTQEP SLTVSPGGT VTLTCGSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPS GTPARFSGSLGGKAAALTL SGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
190	VL of E2M	artificial	aa	QTVVTQEP SLTVSPGGT VTLTCRSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPSGTPARFSG SLGGKAAALTL SGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
191	VH of E2M	artificial	aa	EVQLVESGGGLVQPFGSLKLSCAAASGFTFNKIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KERFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYLSWYAYWGQGTLLVTVSS
192	VH-VL of E2M	artificial	aa	EVQLVESGGGLVQPFGSLKLSCAAASGFTFNKIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KERFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYLSWYAYWGQGTLLVTVSSGGGSGG GGSGGGQTVVTQEP SLTVSPGGT VTLTCRSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPS GTPARFSGSLGGKAAALTL SGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
193	VL-P of E2M	artificial	aa	ELVVTQEP SLTVSPGGT VTLTCRSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPSGTPARFSG SLGGKAAALTL SGVQPEDEAEYICALWYSNRWVFGGGTKLTVL
194	VH-P of E2M	artificial	aa	EVQLLESGGGLVQPFGSLKLSCAAASGFTFNKIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KERFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYLSWYAYWGQGTLLVTVSS
195	VH-VL-P of E2M	artificial	aa	EVQLLESGGGLVQPFGSLKLSCAAASGFTFNKIYAMNWVRQAPGKGLEWVARIRSKYNNIYATYYADSV KERFTI SRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYLSWYAYWGQGTLLVTVSSGGGSGG GGSGGGSELVVTQEP SLTVSPGGT VTLTCRSSTGAVTSGYYPNWVQKPGQAPRGLIGATDMRPS GTPARFSGSLGGKAAALTL SGVQPEDEAEYICALWYSNRWVFGGGTKLTVL

196	VL of F70	artificial	aa	QTVVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
197	VH of F70	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSS
198	VH-VL of F70	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSSGGGSGG GGSGGGSGTAVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAP GTPARFSGSLGGKKAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
199	VL-P of F70	artificial	aa	ELVVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
200	VH-P of F70	artificial	aa	EVQLLESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSS
201	VH-VL-P of F70	artificial	aa	EVQLLESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSSGGGSGG GGSGGGSELVVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAP GTPARFSGSLGGKKAALTLSGVQPEDEAEYYCALWYSNRWVFGGGTKLTVL
202	VL of F12Q	artificial	aa	QTVVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL
203	VH of F12Q	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSS
204	VH-VL of F12Q	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSSGGGSGG GGSGGGSGTAVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAP GTPARFSGSLGGKKAALTLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL
205	VL-P of F12Q	artificial	aa	ELVVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL
206	VH-P of F12Q	artificial	aa	EVQLLESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSS
207	VH-VL-P of F12Q	artificial	aa	EVQLLESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSSGGGSGG GGSGGGSELVVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAP GTPARFSGSLGGKKAALTLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL
208	VL of I2C	artificial	aa	QTVVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAPGTPARFSG SLLGKKAALTLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVL
209	VH of I2C	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSS
210	VH-VL of I2C	artificial	aa	EVQLVESGGGLVQPGGSLKLSCAASGFTFNVIYAMNWVRQAPGKGLEWVARIRSKYNNYATYYADSV KKRFTISRDDSKNTAYLQMNNLKTEDTAVYICVRHGNFGNSYISWWAYWGQGTLLVTVSSGGGSGG GGSGGGSGTAVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGYYPNWVQQKPGQAPRGLIGGTKFLAP

211	VL-P of I2C	artificial	aa	GTPARFSGSLGKKAALTLISGVQPEDEAEYYCVLWYSNRWVFGGTTKLTVL ELVVTFQEPSTLVSPGGTVTLTCGSSGTGAVTSGNYPNWVQKPGQAPRGLIGGTTKFLAPGTPARFSG SLLGKKAALTLISGVQPEDEAEYYCVLWYSNRWVFGGTTKLTVL
212	VH-P of I2C	artificial	aa	EVQLLLESGGLVQPGSLKLSCAAASGFTFNKYAMWVVRQAPGKGLEWVARIRSKYNNYATYYADSV KDRFTIISRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYISYWAYWGQGTTLVTVSS
213	VH-VL-P of I2C	artificial	aa	EVQLLLESGGLVQPGSLKLSCAAASGFTFNKYAMWVVRQAPGKGLEWVARIRSKYNNYATYYADSV KDRFTIISRDDSKNTAYLQMNILKTEDTAVYVCVRHGNFGNSYISYWAYWGQGTTLVTVSSGGSGG GSGGGSELVVTQEPSTLVSPGGTVTLTCGSSGTGAVTSGNYPNWVQKPGQAPRGLIGGTTKFLAP GTPARFSGSLGKKAALTLISGVQPEDEAEYYCVLWYSNRWVFGGTTKLTVL
214	SA21-CD33 E11 HL x I2C HL (His Tag)		nt	CGCTGATTGAAGATATTTGCCGTGCCGCGTGGGGTGCCCTGTGGGAAGATGAT CAGGTGCAGCTGGTGAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAAGTCAAGGTCTCCTGC AAGGTAGCGGGTATACCTTCACAAAAC'TATGGAATGAACTGGGTGAAGCAGGCTCCAGGACAGGGT TTAGAGTGGATGGGCTGGATAAAACACCTACACTGGAGAGCCAACTATGCTGATAAGTTCCAGGGA CGGTTACCATGACTACGGATACCTCTACAGACACTGCC'TATATGGAATACTCCGAACTCCGAGGT GATGACACGGCTGTATATTA'TACTGTGCGCGCTGGAGTTGGAGTGTGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCACTTCGGTCAACCTCTCCTCAGGTGGTGGTGTCTGGCGGCGGGCTCC GGTGGTGGTGGTTC'TGACATCGTATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGGCGAG AGGACCACTCAACTGCAAGTCCAGCCAGAGTGT'TTGAACAGCTCCACGAATAAGAACTCCTTA GCTTGGTACCAGAGAAAACAGGACAGCTCCTAAATTA'CTCTTCC'TGGGCATCTACGCGGGA TCCGGGATCCCTGACCGATTCACTGAGTGGCAGCGGTCTGGGACAGATTTCACTCTCACTATTGACAGC CCGAGCTGAAGATTTGCAACTTACTATTTGTCAAACAGTCTGCCACTTCCCGATCACTCTTTGGC CAAAGGACACGACTGGAGATTAAATCCGAGGTGGTGGTCCGAGGTGAGTGGTGGTGGTGGTGGTGG GGAGGATTGGTGCAGCTGGAGGGTCA'TTGAAACTCTCATGTGCAGCTCTGGATTCACTCTCAAT AAGTACGCCATGAAC'TGGGTCCGACAGCTCCAGGAAAGGT'TTGAAATGGGTGTCTCGCATAGA AGTAAATATAATTA'TATGCAACATATTA'TGCGGATTCAGTGAAGACAGGTTCACTATCCAGA GATGATTCAAAAAACACTGCC'TATCTACAAA'TGAACAACTTGAAAACTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGAACTTCGGTAA'TAGCTACATATCCTTACTGGGCTTACTGGGGCCAAAGG ACTCTGGTCAACCTCTCCTCAGGTGGTGGTGGTTC'TGGCGCGCGCGCTCCGGTGGTGGTGGTTCI CAGACTGTTGTGACTCAGGAACCTTCACTACCGTATCACTGGTGGAAACAGTCACTCACTTGT GGCTCCTCGACTGGGCTGT'TACATCTGGCAACTACCCAACTGGGTCCAAACAAACAGGTCAG GCACCCCGTGGTCTAAATAGGTGGACTAAGTTCCTCGCCCCGCTACTCCTGCAGATTCTCAGGC TCCCTGCTTGGAGGCAAGGCTGCCCTCACTCCTCTCAGGGGTACAGCCAGAGGATGAGGACAGATAT TACTGTGTCTATGTTACAGCAACCGCTGGGTGTTCGGTGGAGAAACCAAACTGACTGTCTCTA CATCACCATCACTAC
215	SA21-CD33 E11 HL x I2C HL H6		aa	RLIEDICLPRWGLWEDDQVQLVQSGAEVKKPESVKYSCKASGYFTTNYGMNWWVKQAPQGLEWM GWINTYTGEPYADKFQGRVTMTTDTSTAYMEIRNLGGDDTAVYYCARWSWDGYVYFYDYWGQ GTSVTVSSGGGGGGGGGGSDIVMTQSPDSLTIVSLGERTTINCKSSQSLDSDSTNKNLSLAWYQ QKPGQPPKILLISWASTRESGIPDRFSGSGSGTFTLTIDSPQEDSATYCYQQSAHFPITFTGQGT LEIKSGGGGSEVQLVESGGGLVQPGGSLKLSCAAASGFTFNKYAMWVVRQAPGKGLEWVARIRSKYN

				NYATYYADSVKDRFTI SRDDSKNTAYLQNMNLKTEDTAVYCYVRHGNFGNSYISYWAYWGQGTIVT VSSGGSGGGSGGGSGQTVVTQEP SLTVSPGGTVTLT CGSSTGAVTSGNYPNWVQOKPGQAPRG LIGGTKFLAPGTPARFSGSLLGGKAAALTLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVLHHHHH H
216	SA25-CD33 E11 HL x I2C HL with His tag		nt	GAGGACATCTGCCGTGCCAGATGGGGCTGCCCTGTGGGAGGACCAGGTGCAGCTGGTGCAGTCTGGA GCTGAGGTGAAGAAGCCTGGAGAGTCAGTCAAGGTCTCTCTGCAAGGCTAGCGGGTATACCTTCACA AACTATGAAATGAAC TGGTGAAGCAGGCTCCAGGACAGGGTTTAGAGTGGATGGGCTGGATAAAC ACCTACACTGGAGAGCCAACCTATGCTGATAAGTTCAGGGACGCGTTACCATGACTACGGATACC TCTACAGCACTGCCATATATGGAATCCGCAACCTCGGAGGTGATGACACGGCTGTATATTACTGT GCGGCTGGAGTTGGAGTGATGTTACTACGTTTACTTTTGTACTACTGGGCCAAGGCACCTTCGGTC ACCGTCTCCTCAGGTGGTGGTTC TGGCGGCGGGCTCCGGTGGTGGTTC TGACATCGTG ATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGAGAGGACCACCATCAACTGCAAGTCC AGCCAGAGTGT TTTAGACAGCTCCACGAATAAGAACTCCTTAGCTTGGTACCAGCAGAAACCCAGGA CAGCCTCCTAAATTACTCCTTTCTTGGGCATCTACGCGGAATCCGGGATCCCTGACCGATTTCAGT GGCAGCGGGTCTGGGACAGATTTCACTCTCACTATTGACAGCCCGCAGCCTGAAGATTCTGCAACT TACTATTGTCAACAGTCTGCCCACTTCCCGATCACCTTTGGCCAAAGGCACACGACTGGAGATTAAA TCCGGAGGTGGTGGCTCCGAGGTGCAGCTGGTGCAGTCTGGAGGAGGATTGGTGCAGCCTGGAGGG TCATTGAAACTCTCATGTGCAGCCTCTTGATTACCTTCAATAAGTACGCCATGAAC TGGTCCGC CAGGCTCCAGGAAAGGGTTTGGAATGGGTGCTCGCATAAGAAGTAAATATAATAATTATGCAACA TATTATGCCGATTCAGTGAAAGACAGGTTCAACCATCTCCAGAGATGATTCAAAAAACACATGCTCTAT CTACAAATGAACAAC TTTGAAAACTGAGGACACTGCCGTGTACTACTGTGTGAGACATGGGAACTTC GGTAATAGTACATATCTACTTGGCTTACTTGGGGCCAAAGGACTCTGGTCACCGTCTCCTCAGGT GGTGGTGGTTC TGGCGGCGGGCTCCGGTGGTGGTTC TACAGACTGTGTGACTCAGGAAACCT TCACTCACCGTATCACCTGGTGGAAACAGTCACTCACTTGTGGCTCTCTCGACTGGGGTGT TACA TCTGGCAACTACCCAAACTGGGTCCAAACAAAACACAGGTACGGCACCCCGTGGTCTAA TAGTGGG ACTAAGTTCTCGCCCCGGTACTCTCTGCCAGATTC TCAGGCTCCCTGCTTGGAGGCAAGGTGCC CTCACCTCTCAGGGGTACAGCCAGAGGATGAGGCAGAAATATTACTGTGTTCTATGTCAGCAAC CGTGGGTGTTCTGGTGGAGGAACCAAACTGACTGTCTCTA CATCACCATCACCATCAC
217	SA25-CD33 E11 HL x I2C HL H6		aa	EDICLPWGCLWEDQVLVQSGAEVKKPGE SVKVSCKASGYTFTNYGMWVVKQAPGQGLEWMGWIN TYTGEPTYADKFQGRVTMTTDTSTAYMEIRNLGDDDTAVYCARWSWSDGYVYFYFDYWQGTSV TVSSGGSGGGSGGGSGD IVMTQSPDSLTVSLGERTTINCKSSQSVLDSSTNKNLSLAWYQQKPG QPPKLLLSWASTRESGIPDRFSGSGSGTDFTLTIDSQPEDSATYCYQSAHFPITTFGQGTGLEIK SGGGGSEVQLVE SGGGLVQPGSLKLSCAAASGFTFNKYAMNWVRQAPGKGLEWVARIRSKYNNYAT YYADSVKDRFTI SRDDSKNTAYLQNMNLKTEDTAVYCYVRHGNFGNSYISYWAYWGQGTIVTVSSG GGSGGGSGGGSGQTVVTQEP SLTVSPGGTVTLT CGSSTGAVTSGNYPNWVQOKPGQAPRGLIGG TKFLAPGTPARFSGSLLGGKAAALTLSGVQPEDEAEYYCVLWYSNRWVFGGGTKLTVLHHHHH
218	SA08-CD33 E11 HL x I2C HL with His tag		nt	CAGGGCCTGATCGGCGACATCTGCCCTGCCAGATGGGGCTGCCCTGTGGGCGACTCCCGTGAACACAG GTGCAGCTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCCTGCAAG GCTAGCGGGTATACCTTCACAAACTATGGAATGAAC TGGGTGAAGCAGCGCTCCAGGACACAGGGTTTA

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226	CD33 E11 HL x I2C HL (His Tag)	nt	QGLIGDICI LPRWGCLWGDVVKVQVLVQSGAEVKKPGEVSVKVSCKASGYTFN YGMNWNVVKQAPGQGL EWMGWINTYTGEPTYADKFKQGRVTMTTDTSTAYMEIRNLGGDDTAVYICARWSWSDGYVYVFDY WGQGTSVTVSSGGGGGGGGSDIVMTQSPDSLTVSLGERTTINCKSSQSVLSDSTNKNLSLA WYQQKPGQPPKLLLSWASTRELSGIPDRFSGSGGTDFLTIDSPQPEDSATYCYQQSAHFPIITFGQ GTRLEIKSGGGSEVQLVLESGGGLVQPGSLKLSCAASGFTFNKYAMNWNVRQAPGKGLEWVARIRS KYNNTATYYADSVKDRFTISRDDSKNTAYLQMNLLKTEDTAVYCVVRHGNFNSYISYWAYWGQGT LIVTVSSGGGGGGGGGSGTAVVTEPSTLTVSPGGFTLTTCGSSGTGAFTSGNYPNWVQOKPGQA PRGLIGGTFKFLAPGTPARFSGSLGGKKAALTLISGVQVPEDEAEYYCVLWVSNRWVFGGGTKLTVL CAGGTGACGTGGTGCAGTCTGGAGCTGAGGTGAAGAAAGCCTGGAGAGTCAGTCAAGGTCTCTCTGC AAGGTAGCGGGTATACCTTTCACAAAATATGGAAATGAACTGGGTGAAGCAGGTCCAGGACAGGGT TTAGAGTGGATGGGTGGATAAACACCTACACTGGAGAGCCAACTATGCTGATAAAGTTCAGGGA CGGTACCATGACTACGGATACCTCTACCCAGCACTGCCATATATGGAAAATCCGCAACCTCGGAGGT GATGACACGGCTGTATATTAATGTGCGCGCTGGAGTTGGAGTGAATGTTACTACGTTTACTTTGAC TACTGGGGCCAAAGGCATTCGGTCAACCGTCTCCTCAGGTGGTGGTGTCTGGCGCGCGCGGTCC GGTGGTGGTGTCTGACATCGTGATGACACAGTCTCCAGACTCCCTGACTGTGTCTCTGGCGGAG AGGACCACCATCAACTGCAAGTCCAGCCAGAGTGTATAGACAGCTCCACGAATAAGAACTCCTTA GCTTGGTACCAGAGAAAACAGGACAGCTCCTAAAATTACTCTCTTCTGGGCATCTACGCGGAA TCCGGGATCCCTGACCGATTCAAGTGGCAGCGGTCTGGGACAGATTTCACCTCACTATTGACAGC CCGACGCTGAAGATTCTGCAACTTACTATTTGTCAAACAGTCTGCCACTTCCCGATCACTTTGGC CAAGGACACGACTGGAGATTAAATCCGGAGGTGGTGGTCCAGAGTGCAGTGGTCGAGTCTGGA GGAGGATTGGTGACGCTGGAGGGTCAATTGAAACTCTCATGTGCAGCTCTGGATTCACTTCAAT AAGTACGCCATGAAC TGGGTCCCGCAGGTCCAGGAAAAGGT TGGAA TGGGTGCTCGCATAGA AGTAAAATATAATTAATGCAACATATTAATGCCGATTTCAGTGAAGACAGAGTTTACCATCTCCAGA GATGATTCAAAAACACTGCCATCTACAAAATGAACAAC TTGAAAACTTGAGGACACTGCCGTGTAC TACTGTGTGAGACATGGGAACTTCGGTAATAGCTACATATCTTACTGGGCTTACTGGGCGCAAGGG ACTCTGGTCACCGTCTCCTCAGGTGGTGGTGTCTGGCGCGCGCGGTCCCGTGGTGGTGTCT CAGACTGTGTGACTCAGGAACCTTCACTACCGTATCACTGGTGAACAGTCACACTCACTGT GGCTCCTCGACTGGGCTGTACATCTGGCAACTACCCAACTGGGTCCAAACAAAAACAGGTCAAG GCACCCCGTGGTCTTAATAGGTGGGACTAAAGTTCCCTCGCCCCCGGTACTCTTCGACAGATTCTCAGGC TCCCTGTCTGGAGGCAAGGTGCCCTCACTCTCAGGGGTACAGCAGAGGATGAGGACAGAAATAT TACTGTGTCTTATGTTACAGCAACCGCTGGGTGTCTCGGTGGAGGAACCAAACTGACTGTCTCTA CATCACCATCACCATCAC
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Claims

1. A pharmaceutical composition comprising a CD33 targeting compound and at least one epigenetic factor, wherein
 - (a) the CD33 targeting compound is a bispecific single chain antibody construct comprising a first binding domain specifically binding to CD33 and a second binding domain specifically binding to CD3; and
 - (b) the at least one epigenetic factor is selected from the group consisting of histone deacetylase (HDAC) inhibitor panobinostat, DNA methyltransferase (DNMT) I inhibitor 5-azacitidine, hydroxyurea, and Granulocyte-Colony Stimulating Factor (G-CSF), and wherein
 - (i) the epigenetic factor is administered prior to the administration of the CD33 targeting compound;
 - (ii) the epigenetic factor is administered subsequent to the administration of the CD33 targeting compound; or
 - (iii) the epigenetic factor and the CD33 targeting compound are administered simultaneously.
2. The pharmaceutical composition of claim 1, wherein a first dose of the epigenetic factor is administered prior to the start of the administration of the CD33 targeting compound.
3. The pharmaceutical composition of claim 2, wherein the administration of the epigenetic factor is continued during the administration of the CD33 targeting compound.
4. The pharmaceutical composition of any one of claims 1-3, wherein the bispecific single chain antibody construct binds to human and cynomolgous CD3 and human and cynomolgous CD33.
5. The pharmaceutical composition of any one of claims 1-4, wherein the bispecific single chain antibody construct comprises
 - a first binding domain specifically binding to CD33 and comprising a VL chain having an amino acid sequence selected from the group consisting of SEQ ID NO: 6, 24, 42, 60, 78, 96, 114 and 132 and a VH chain having

- an amino acid sequence selected from the group consisting of SEQ ID NO:1, 19, 37, 55, 73, 91, 109 and 127; and
- a second binding domain specifically binding to CD3 and comprising a VL chain having an amino acid sequence selected from the group consisting of SEQ ID NO: 154, 157, 160, 163, 166, 169 and 172 and a VH chain having an amino acid sequence selected from the group consisting of SEQ ID NO: 155, 158, 161, 164, 167, 170 and 173.
6. The pharmaceutical composition of claim 5, wherein the bispecific single chain antibody construct comprises an amino acid sequence selected from the group consisting of SEQ ID NOs: 13, 15, 17, 31, 33, 35, 49, 51, 53, 67, 69, 71, 85, 87, 89, 103, 105, 107, 121, 123, 125, 139, 141, 143, 148, 150, 152, 215, 217, 219, 221, 223, 225 and 227.
 7. The pharmaceutical composition of any one of claims 1 to 6 when used in the treatment of a myeloid leukemia.
 8. The pharmaceutical composition of claim 7, wherein the myeloid leukemia is selected from the group consisting of acute myeloblastic leukemia, chronic neutrophilic leukemia, myeloid dendritic cell leukemia, accelerated phase chronic myelogenous leukemia, acute myelomonocytic leukemia, juvenile myelomonocytic leukemia, chronic myelomonocytic leukemia, acute basophilic leukemia, acute eosinophilic leukemia, chronic eosinophilic leukemia, Acute megakaryoblastic leukemia, essential thrombocytosis, acute erythroid leukemia, polycythemia vera, myelodysplastic syndrome, acute panmyelosis, myeloid sarcoma, and acute biphenotypic leukaemia.
 9. Use of an epigenetic factor in the manufacture of an agent in the amelioration and/or treatment of a myeloid leukemia, wherein the epigenetic factor increases the responsiveness of a patient to a CD33 targeting compound, wherein
 - (a) the CD33 targeting compound is a bispecific single chain antibody construct comprising a first binding domain specifically binding to CD33 and a second binding domain specifically binding to CD3; and
 - (b) the at least one epigenetic factor is selected from the group consisting of histone deacetylase (HDAC) inhibitor panobinostat, DNA

methyltransferase (DNMT) I inhibitor 5-azacitidine, hydroxyurea, and Granulocyte-Colony Stimulating Factor (G-CSF).

10. A method of treatment of a myeloid leukemia comprising administering to a subject in need thereof a pharmaceutical composition according to any one of claims 1-8.
11. The method according to claim 10, wherein the myeloid leukemia is selected from the group consisting of acute myeloblastic leukemia, chronic neutrophilic leukemia, myeloid dendritic cell leukemia, accelerated phase chronic myelogenous leukemia, acute myelomonocytic leukemia, juvenile myelomonocytic leukemia, chronic myelomonocytic leukemia, acute basophilic leukemia, acute eosinophilic leukemia, chronic eosinophilic leukemia, Acute megakaryoblastic leukemia, essential thrombocytosis, acute erythroid leukemia, polycythemia vera, myelodysplastic syndrome, acute panmyelosis, myeloid sarcoma, and acute biphenotypic leukaemia.
12. A kit when used in the amelioration and/or treatment of a myeloid leukemia, comprising a pharmaceutical composition of any one of claims 1 to 8.

Figure 1

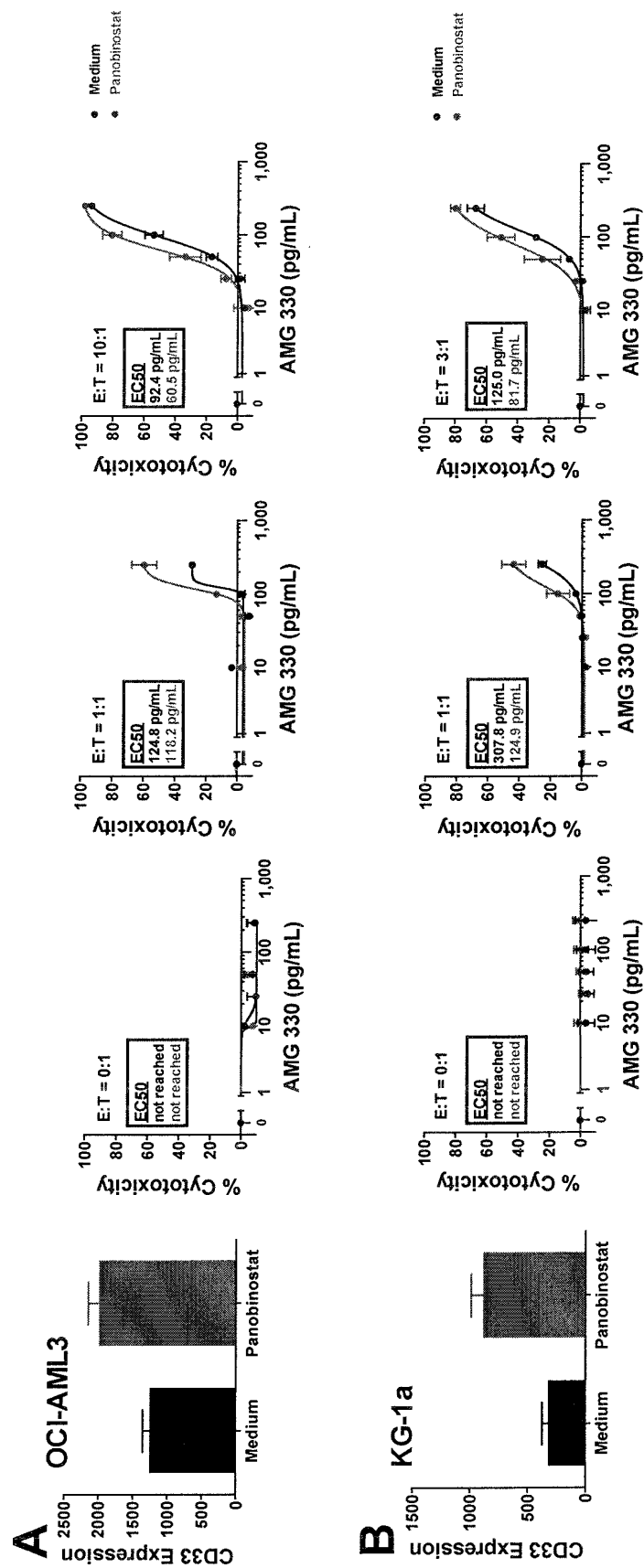


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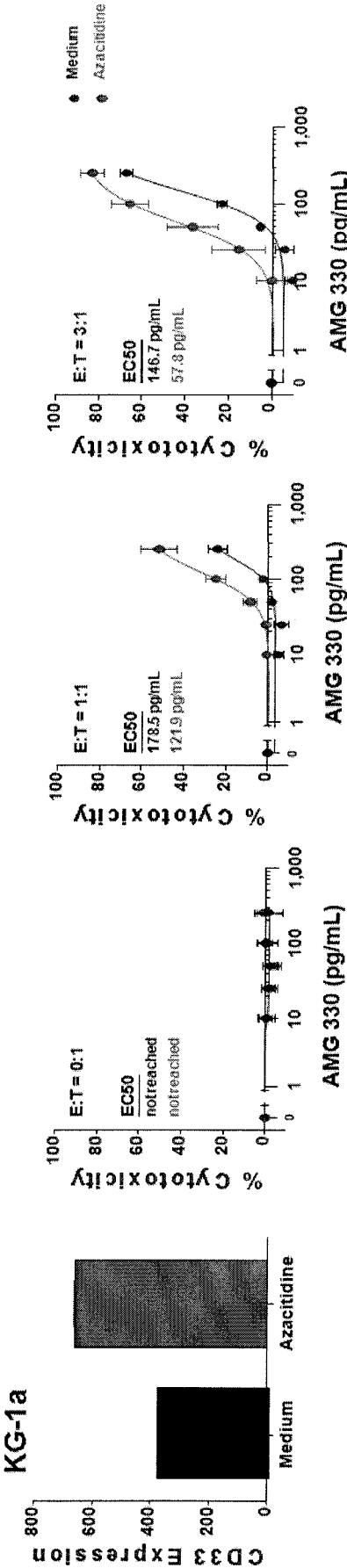
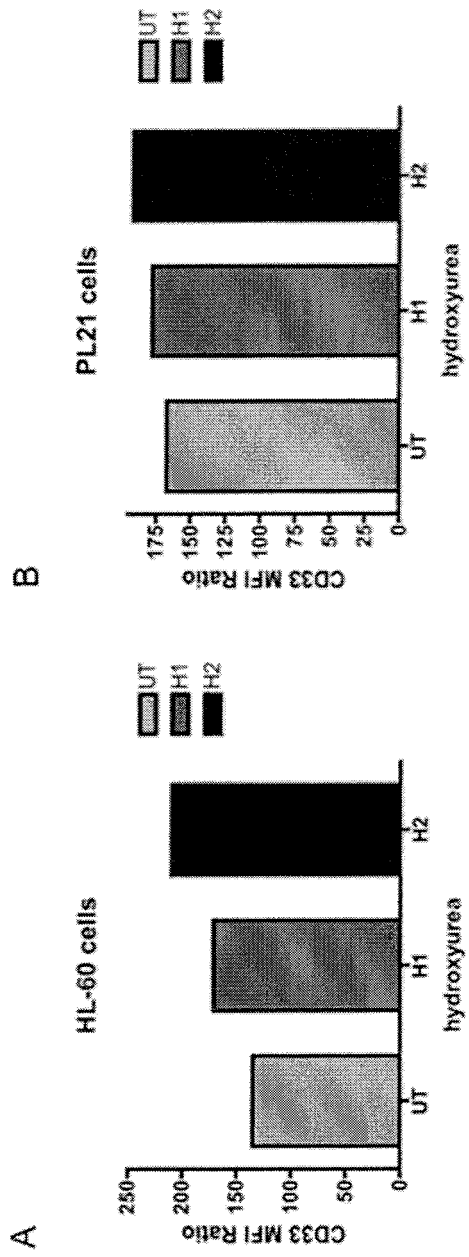


Figure 3



4/5

Figure 4

Patient 2

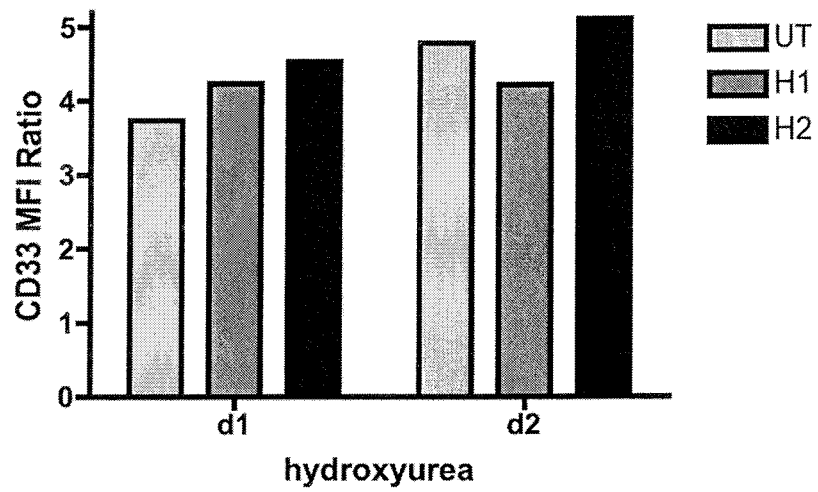


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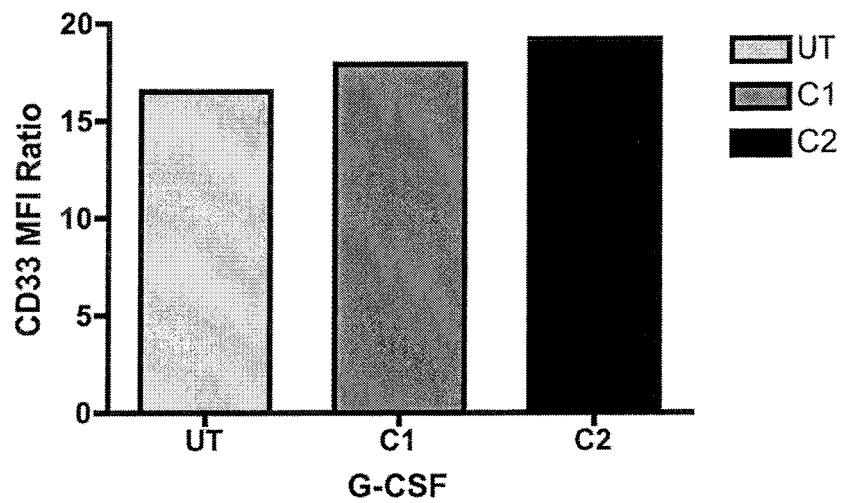
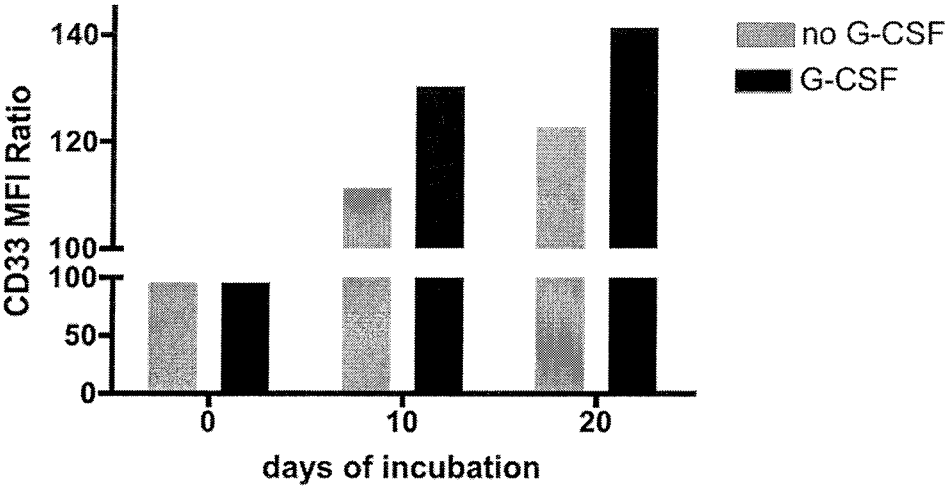
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Figure 6

Patient 1



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Amgen Inc.

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Glu Arg Thr Thr Ile Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser
 20 25 30

Ser Lys Asn Lys Asn Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln
 35 40 45

Pro Pro Lys Leu Leu Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile
 50 55 60

Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr
 65 70 75 80

Ile Asp Ser Leu Gln Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln
 85 90 95

Ser Ala His Phe Pro Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile
 100 105 110

Lys

<210> 7
 <211> 17
 <212> PRT
 <213> artificial

<220>
 <223> CD33 LCDR1 of AH3

<400> 7

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
 1 5 10 15

Ala

eof-seq1.txt

<210> 8
 <211> 7
 <212> PRT
 <213> artificial

<220>
 <223> CD33 LCDR2 of AH3

<400> 8

Trp Ala Ser Thr Arg Glu Ser
 1 5

<210> 9
 <211> 9
 <212> PRT
 <213> artificial

<220>
 <223> CD33 LCDR3 of AH3

<400> 9

Gln Gln Ser Ala His Phe Pro Ile Thr
 1 5

<210> 10
 <211> 339
 <212> DNA
 <213> artificial

<220>
 <223> CD33 VL of AH3

<400> 10
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 atcaactgca agtccagcca gagtgtttta gacagctcca agaataagaa ctccttagct 120
 tggtagcagc agaaaccagg acagcctcct aaattactcc tttcctgggc atctacgcgg 180
 gaatccggga tccctgaccg attcagtggc agcgggtctg ggacagattt cactctcact 240
 attgacagcc tgcagcctga agattctgca acttactatt gtcaacagtc tgcccacttc 300
 ccgatcacct ttggccaagg gacacgactg gagattaaa 339

<210> 11
 <211> 250
 <212> PRT
 <213> artificial

<220>
 <223> CD33 CD33 HL of AH3

<400> 11

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15

eo1f-seq1.txt

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

Gly Met Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
50 55 60

Lys Gly Arg Val Thr Met Ser Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Leu Glu Ile Asn Ser Leu Arg Ser Asp Asp Thr Ala Ile Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys
245 250

<210> 12
<211> 750

eolf-seq1.txt

<212> DNA
<213> artificial

<220>
<223> CD33 HL of AH3

<400> 12
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tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaggcaggct 120
ccaggacagg gtttagagtg gatgggctgg ataaacacct acactggaga gccaacatat 180
gctgatgact tcaagggacg ggttaccatg tcttcggata cctctaccag cactgcctat 240
ttggaaatca acagcctcag aagtgatgac acggctatat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcaggtg gtgggtggtc tggcggcggc ggctccggtg gtgggtggtc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa 750

<210> 13
<211> 505
<212> PRT
<213> artificial

<220>
<223> CD33 AH3 HL x H2C HL

<400> 13
Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1 5 10 15
Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30
Gly Met Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45
Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
50 55 60
Lys Gly Arg Val Thr Met Ser Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

eo1f-seq1.txt

Leu Glu Ile Asn Ser₈₅ Leu Arg Ser Asp₉₀ Thr Ala Ile Tyr Tyr Cys
 Ala Arg Trp Ser₁₀₀ Trp Ser Asp Gly Tyr₁₀₅ Tyr Val Tyr Phe Asp₁₁₀ Tyr Trp
 Gly Gln Gly₁₁₅ Thr Thr Val Thr Val₁₂₀ Ser Ser Gly Gly Gly₁₂₅ Gly Ser Gly
 Gly Gly₁₃₀ Gly Ser Gly Gly Gly₁₃₅ Gly Ser Asp Ile Val₁₄₀ Met Thr Gln Ser
 Pro Asp Ser Leu Thr Val₁₅₀ Ser Leu Gly Glu Arg₁₅₅ Thr Thr Ile Asn Cys₁₆₀
 Lys Ser Ser Gln Ser₁₆₅ Val Leu Asp Ser Ser₁₇₀ Lys Asn Lys Asn Ser₁₇₅ Leu
 Ala Trp Tyr Gln Gln Lys Pro Gly Gln₁₈₅ Pro Pro Lys Leu Leu₁₉₀ Leu Ser
 Trp Ala Ser₁₉₅ Thr Arg Glu Ser Gly₂₀₀ Ile Pro Asp Arg Phe₂₀₅ Ser Gly Ser
 Gly Ser₂₁₀ Gly Thr Asp Phe Thr₂₁₅ Leu Thr Ile Asp Ser₂₂₀ Leu Gln Pro Glu
 Asp Ser Ala Thr Tyr Tyr₂₃₀ Cys Gln Gln Ser Ala₂₃₅ His Phe Pro Ile Thr₂₄₀
 Phe Gly Gln Gly Thr₂₄₅ Arg Leu Glu Ile Lys₂₅₀ Ser Gly Gly Gly Gly₂₅₅ Ser
 Glu Val Gln Leu Val Glu Ser Gly Gly₂₆₅ Gly Leu Val Gln Pro₂₇₀ Gly Gly
 Ser Leu Lys₂₇₅ Leu Ser Cys Ala Ala₂₈₀ Ser Gly Phe Thr Phe₂₈₅ Asn Lys Tyr
 Ala Met₂₉₀ Asn Trp Val Arg Gln₂₉₅ Ala Pro Gly Lys Gly₃₀₀ Leu Glu Trp Val
 Ala Arg Ile Arg Ser Lys₃₁₀ Tyr Asn Asn Tyr Ala₃₁₅ Thr Tyr Tyr Ala Asp₃₂₀
 Ser Val Lys Asp Arg₃₂₅ Phe Thr Ile Ser Arg₃₃₀ Asp Asp Ser Lys Asn₃₃₅ Thr

eo1f-seq1.txt

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 14
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AH3 HL x H2C HL

<400> 14
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tcctgcaagg ctacgaggta taccttcaca aactatggaa tgaactgggt gaggcaggct 120
ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
gctgatgact tcaagggacg gggtaccatg tcttcggata cctctaccag cactgcctat 240

eolf-seq1.txt

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tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcaggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtcacagc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg 780
gtcgagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc 840
tctggattca ctttcaataa gtacgccatg aactgggtcc gccagggtcc aggaaagggt 900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaaa acaggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
ggtaatagct acatatccta ctgggcttac tggggccaag ggactctggt caccgtctcc 1140
tcagggtgtg gtggttctgg cggcggcggc tccggtggtg gtggttctca gactgttgtg 1200
actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtggctcc 1260
tcgactgggg ctgttacatc tggctactac ccaaactggg tccaacaaaa accaggtcag 1320
gcaccccggt gtctaatagg tgggactaag ttctctgccc ccggtactcc tgccagattc 1380
tcaggctccc tgcttggagg caaggctgcc ctcaccctct caggggtaca gccagaggat 1440
gaggcagaat attactgtgc tctatggtac agcaaccgct ggggtgttcgg tggaggaacc 1500
aaactgactg tccta 1515

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<210> 15
<211> 505
<212> PRT
<213> artificial

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```

<220>
<223> CD33 AH3 HL x F12Q HL

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```

<400> 15

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Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1           5           10          15

```

```

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
          20          25          30

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eo1f-seq1.txt

Gly Met Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
50 55 60

Lys Gly Arg Val Thr Met Ser Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Leu Glu Ile Asn Ser Leu Arg Ser Asp Asp Thr Ala Ile Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
275 280 285

eo1f-seq1.txt

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 305 310 315 320

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
 355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
 385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
 420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
 435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
 450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
 465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
 485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
 500 505

<210> 16
 <211> 1515
 <212> DNA
 <213> artificial

<220>

<223> CD33 AH3 HL x F12Q HL

<400> 16

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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacatat      180
gctgatgact tcaagggacg ggttaccatg tcttcggata cctctaccag cactgcctat      240
ttggaaatca acagcctcag aagtgatgac acggctatat attactgtgc gcgctggagt      300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc      360
tcctcagggtg gtggtgggttc tggcggcggc ggctccgggtg gtggtgggttc tgacatcgtg      420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc      480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag      540
cagaaaccag gacagcctcc taaattactc ctttctctggg catctacgcg ggaatccggg      600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc      660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc      720
tttggccaaag ggacacgact ggagattaaa tccggagggtg gtgggtccga ggtgcagctg      780
gtcgagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaaactctc atgtgcagcc      840
tctggattca ccttcaatag ctacgccatg aactgggtcc gccaggctcc aggaaagggt      900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat      960
tcagtgaaaag gcagggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa     1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc     1080
ggtaatagct acgtttcctg gtgggcttac tggggccaag ggactctggt caccgtctcc     1140
tcaggtggtg gtggttcttg cggcggcggc tccggtggtg gtggttctca gactgttgtg     1200
actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtggctcc     1260
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tcaggctccc tgcttgagg caaggctgcc ctaccctct caggggtaca gccagaggat     1440
gaggcagaat attactgtgt tctatggtac agcaaccgct ggggtgttcgg tggaggaacc     1500
aaactgactg tccta                                             1515

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<210> 17

<211> 505

<212> PRT

<213> artificial

<220>

<223> CD33 AH3 HL x I2C HL

eo1f-seq1.txt

<400> 17

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Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1      5      10     15
Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20     25     30
Gly Met Asn Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35     40     45
Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
50     55     60
Lys Gly Arg Val Thr Met Ser Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65     70     75     80
Leu Glu Ile Asn Ser Leu Arg Ser Asp Asp Thr Ala Ile Tyr Tyr Cys
85     90     95
Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100    105    110
Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115    120    125
Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130    135    140
Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145    150    155    160
Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165    170    175
Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180    185    190
Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195    200    205
Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210    215    220
Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225    230    235    240

```

eo1f-seq1.txt

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

eof-seq1.txt

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 18
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AH3 HL x I2C HL

<400> 18
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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
gctgatgact tcaagggacg ggttaccatg tcttcggata cctctaccag cactgcctat 240
ttgaaaatca acagcctcag aagtgatgac acggctatat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcagggtg gtggtgggtc tggcggcggc ggctccgggtg gtggtgggtc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactctac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg 780
gtcagagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc 840
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tcaggctccc tgcttggagg caaggctgcc ctcaccctct caggggtaca gccagaggat 1440
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aaactgactg tccta

1515

<210> 19
 <211> 122
 <212> PRT
 <213> artificial

<220>
 <223> CD33 VH of AF5

<400> 19

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 Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
 50 55 60

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80

Leu Glu Leu His Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser
 115 120

<210> 20
 <211> 5
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HCDR1 of AF5

<400> 20

Asn Tyr Gly Met Asn
 1 5

<210> 21
 <211> 17
 <212> PRT

<213> artificial

<220>

<223> CD33 HCDR2 of AF5

<400> 21

Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe Lys
1 5 10 15

Gly

<210> 22

<211> 13

<212> PRT

<213> artificial

<220>

<223> CD33 HCDR3 of AF5

<400> 22

Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr
1 5 10

<210> 23

<211> 366

<212> DNA

<213> artificial

<220>

<223> CD33 VH of AF5

<400> 23

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ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat	180
gctgatgact tcaagggacg gggtaccatg acttcggata cctctaccag cactgcctat	240
ttggaactcc acaacctcag aagtgatgac acggctgtat attactgtgc gcgctggagt	300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc	360
tcctca	366

<210> 24

<211> 113

<212> PRT

<213> artificial

<220>

<223> CD33 VL of AF5

<400> 24

eof-seq1.txt

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1 5 10 15

Glu Arg Thr Thr Ile Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser
20 25 30

Ser Lys Asn Lys Asn Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln
35 40 45

Pro Pro Lys Leu Leu Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile
50 55 60

Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr
65 70 75 80

Ile Asp Ser Leu Gln Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln
85 90 95

Ser Ala His Phe Pro Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile
100 105 110

Lys

<210> 25
<211> 17
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR1 of AF5
<400> 25

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
1 5 10 15

Ala

<210> 26
<211> 7
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR2 of AF5
<400> 26

Trp Ala Ser Thr Arg Glu Ser
1 5

eof-seq1.txt

<210> 27
 <211> 9
 <212> PRT
 <213> artificial

<220>
 <223> CD33 LCDR3 of AF5

<400> 27

Gln Gln Ser Ala His Phe Pro Ile Thr
 1 5

<210> 28
 <211> 339
 <212> DNA
 <213> artificial

<220>
 <223> CD33VL of AF5

<400> 28
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 tgggtaccagc agaaaccagg acagcctcct aaattactcc tttcctgggc atctacgcgg 180
 gaatccggga tccctgaccg attcagtggc agcgggtctg ggacagattt cactctcact 240
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<210> 29
 <211> 250
 <212> PRT
 <213> artificial

<220>
 <223> CD33HL of AF5

<400> 29

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 Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
 50 55 60

eo1f-seq1.txt

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Leu Glu Leu His Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys
245 250

<210> 30
<211> 750
<212> DNA
<213> artificial

<220>
<223> CD33HL of AF5

<400> 30
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tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct 120
ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat 180

eof-seq1.txt

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tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcaggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtcacgcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa 750

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<210> 31
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 AF5 HL x H2C HL

<400> 31

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 Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
 50 55 60

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80

Leu Glu Leu His Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125

eo1f-seq1.txt

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

eof-seq1.txt

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 32
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AF5 HL x H2C HL

<400> 32
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ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
gctgatgact tcaagggacg ggttaccatg acttcggata cctctaccag cactgcctat 240
ttggaactcc acaacctcag aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcaggtg gtgggtggtc tggcggcggc ggctccggtg gtgggtggtc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
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eof-seq1.txt

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gtcagagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc 840
tctggattca ccttcaataa gtacgccatg aactgggtcc gccagggtcc aggaaagggt 900
ttggaatggg ttgctcgcac aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaaaag acaggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
ggtaatagct acatatccta ctgggcttac tggggccaag ggactctggt caccgtctcc 1140
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tcgactgggg ctgttacatc tggctactac ccaaactggg tccaacaaaa accaggtcag 1320
gcaccccgtg gtctaataagg tgggactaag ttcctcgccc ccggtactcc tgccagattc 1380
tcaggctccc tgcttgagg caaggctgcc ctcacctct caggggtaca gccagaggat 1440
gaggcagaat attactgtgc tctatggtac agcaaccgct ggggtgttcgg tggaggaacc 1500
aaactgactg tccta 1515

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<210> 33
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 AF5 HL x F12Q HL

<400> 33

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 Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
 50 55 60

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80

Leu Glu Leu His Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys

Ala Arg Trp Ser₁₀₀ Trp Ser Asp Gly Tyr₁₀₅ Tyr Val Tyr Phe Asp₁₁₀ Tyr Trp
Gly Gln Gly₁₁₅ Thr Thr Val Thr Val₁₂₀ Ser Ser Gly Gly₁₂₅ Gly Ser Gly
Gly Gly₁₃₀ Gly Ser Gly Gly Gly₁₃₅ Gly Ser Asp Ile Val₁₄₀ Met Thr Gln Ser
Pro Asp Ser Leu Thr Val₁₅₀ Ser Leu Gly Glu Arg₁₅₅ Thr Thr Ile Asn Cys₁₆₀
Lys Ser Ser Gln Ser₁₆₅ Val Leu Asp Ser Ser₁₇₀ Lys Asn Lys Asn Ser₁₇₅ Leu
Ala Trp Tyr Gln₁₈₀ Gln Lys Pro Gly Gln₁₈₅ Pro Pro Lys Leu Leu₁₉₀ Leu Ser
Trp Ala Ser₁₉₅ Thr Arg Glu Ser Gly₂₀₀ Ile Pro Asp Arg Phe₂₀₅ Ser Gly Ser
Gly Ser₂₁₀ Gly Thr Asp Phe Thr₂₁₅ Leu Thr Ile Asp Ser₂₂₀ Leu Gln Pro Glu
Asp Ser₂₂₅ Ala Thr Tyr Tyr₂₃₀ Cys Gln Gln Ser Ala₂₃₅ His Phe Pro Ile Thr₂₄₀
Phe Gly Gln Gly Thr₂₄₅ Arg Leu Glu Ile Lys₂₅₀ Ser Gly Gly Gly₂₅₅ Gly Ser
Glu Val Gln Leu Val₂₆₀ Glu Ser Gly Gly₂₆₅ Gly Leu Val Gln Pro₂₇₀ Gly Gly
Ser Leu Lys₂₇₅ Leu Ser Cys Ala Ala₂₈₀ Ser Gly Phe Thr Phe₂₈₅ Asn Ser Tyr
Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly₃₀₀ Leu Glu Trp Val
Ala Arg Ile Arg Ser Lys₃₁₀ Tyr Asn Asn Tyr Ala₃₁₅ Thr Tyr Tyr Ala Asp₃₂₀
Ser Val Lys Gly Arg₃₂₅ Phe Thr Ile Ser Arg₃₃₀ Asp Asp Ser Lys Asn₃₃₅ Thr

eo1f-seq1.txt

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 34
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AF5 HL x F12Q HL

<400> 34
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ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
gctgatgact tcaagggacg ggttaccatg acttcggata cctctaccag cactgcctat 240
ttggaactcc acaacctcag aagtgatgac acggctgtat attactgtgc gcgctggagt 300

eof-seq1.txt

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tcctcagggtg gtggtggttc tggcggcggc ggctccgggtg gtggtggttc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc 660
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tttggccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg 780
gtcgagtctg gaggaggatt ggtgcagcct ggagggcat tgaaactctc atgtgcagcc 840
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tcagtgaag gcagggtcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
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tcaggctccc tgcttggagg caaggctgcc ctcaccctct caggggtaca gccagaggat 1440
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aaactgactg tccta 1515

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<210> 35
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 AF5 HL x I2C HL
 <400> 35

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 Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45

eo1f-seq1.txt

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
50 55 60

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Leu Glu Leu His Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val

eo1f-seq1.txt
300

290

295

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 36
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AF5 HL x I2C HL

eof-seq1.txt

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<400> 36
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ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
gctgatgact tcaagggacg ggttaccatg acttcggata cctctaccag cactgcctat 240
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tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcagggtg gtgggtggtc tggcggcggc ggctccgggtg gtgggtggtc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
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aaactgactg tccta 1515

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<210> 37
<211> 122
<212> PRT
<213> artificial

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<220>
<223> CD33 VH of AC8

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<400> 37

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eo1f-seq1.txt

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 20 25 30
 Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45
 Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp 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 Ile Arg Asn Leu Arg Asn Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95
 Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110
 Gly Gln Gly Thr Thr Val Thr Val Ser Ser
 115 120

<210> 38
 <211> 5
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HCDR1 of AC8

<400> 38

Asn Tyr Gly Met Asn
 1 5

<210> 39
 <211> 17
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HCDR2 of AC8

<400> 39

Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe Lys
 1 5 10 15

Gly

eo1f-seq1.txt

<210> 40
 <211> 13
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HCDR3 of AC8

<400> 40

Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr
 1 5 10

<210> 41
 <211> 366
 <212> DNA
 <213> artificial

<220>
 <223> CD33 VH of AC8

<400> 41
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 ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
 gctgatgact tcaagggacg ggttaccatg actacggata cctctaccag cactgcctat 240
 atggaaaatcc gcaacctcag aaatgatgac acggctgtat attactgtgc gcgctggagt 300
 tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
 tcctca 366

<210> 42
 <211> 113
 <212> PRT
 <213> artificial

<220>
 <223> CD33 VL of AC8

<400> 42

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu Thr Val Ser Leu Gly
 1 5 10 15

Glu Arg Thr Thr Ile Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser
 20 25 30

Ser Lys Asn Lys Asn Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln
 35 40 45

Pro Pro Lys Leu Leu Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile

50

55

Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr
65 70 75 80

Ile Asp Ser Leu Gln Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln
85 90 95

Ser Ala His Phe Pro Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile
100 105 110

Lys

<210> 43
<211> 17
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR1 of AC8

<400> 43

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
1 5 10 15

Ala

<210> 44
<211> 7
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR2 of AC8

<400> 44

Trp Ala Ser Thr Arg Glu Ser
1 5

<210> 45
<211> 9
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR3 of AC8

<400> 45

Gln Gln Ser Ala His Phe Pro Ile Thr
1 5

eof-seq1.txt

<210> 46
 <211> 339
 <212> DNA
 <213> artificial

<220>
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<400> 46
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 tgggtaccagc agaaaccagg acagcctcct aaattactcc tttcctgggc atctacgcgg 180
 gaatccggga tccctgaccg attcagtggc agcgggtctg ggacagattt cactctcact 240
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<210> 47
 <211> 250
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HL of AC8

<400> 47

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
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Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp 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 Ile Arg Asn Leu Arg Asn Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly

115

120

125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
 130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
 145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
 165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
 180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
 195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
 210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
 225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys
 245 250

<210> 48
 <211> 750
 <212> DNA
 <213> artificial

<220>
 <223> CD33 HL of AC8

<400> 48
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 ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
 gctgatgact tcaagggacg gggtaccatg actacggata cctctaccag cactgcctat 240
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 tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
 tcctcagggtg gtggtgggtc tggcggcggc ggctccggtg gtggtgggtc tgacatcgtg 420
 atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
 aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
 cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600

eof-seq1.txt

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 ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
 tttggccaag ggacacgact ggagattaaa 750

<210> 49
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 AC8 HL x H2C HL

<400> 49

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp 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 Ile Arg Asn Leu Arg Asn Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
 130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
 145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
 165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser

180

185

190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
 195 200 205
 Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
 210 215 220
 Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
 225 230 235 240
 Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
 245 250 255
 Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 260 265 270
 Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
 275 280 285
 Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 290 295 300
 Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 305 310 315 320
 Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 325 330 335
 Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 340 345 350
 Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
 355 360 365
 Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 370 375 380
 Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
 385 390 395 400
 Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 405 410 415
 Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
 420 425 430

eof-seq1.txt

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 50
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AC8 HL x H2C HL

<400> 50
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ccaggacagg gtttaaagtg gatgggctgg ataaacacct acactggaga gccaacatat 180
gctgatgact tcaaggggacg gggtaccatg actacggata cctctaccag cactgcctat 240
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tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
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atgacacagt ctccagactc cctgactgtg tctctgggag agaggaccac catcaactgc 480
aagtcagacc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
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ttggaatggg ttgctcgcag aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaaag acagggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020

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<210> 51
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 AC8 HL x F12Q HL

<400> 51

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp 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 Ile Arg Asn Leu Arg Asn Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
 130 135 140

eo1f-seq1.txt

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val

385 390 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 52
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AC8 HL x F12Q HL

<400> 52
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gctgatgact tcaagggacg ggttaccatg actacggata cctctaccag cactgcctat 240
atggaaatcc gcaacctcag aaatgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcagggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtcacagg agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tctctctcac tattgacagc 660
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eof-seq1.txt

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tctggattca ctttcaatag ctacgccatg aactgggtcc gccagggtcc aggaaaggg 900
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tcagtgaag gcaggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
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aaactgactg tccta 1515

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<210> 53
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 AC8 HL x I2C HL

<400> 53

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp 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 Ile Arg Asn Leu Arg Asn Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

eo1f-seq1.txt

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

eo1f-seq1.txt

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 54
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AC8 HL x I2C HL

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gctgatgact tcaagggacg ggttaccatg actacggata cctctaccag cactgcctat 240
atggaaaatcc gcaacctcag aaatgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgctc 360

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cagaaaccag	gacagcctcc	taaattactc	ctttcctggg	catctacgcg	ggaatccggg	600
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<211>	122
<212>	PRT
<213>	artificial
<220>	
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<400>	55

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

eof-seq1.txt

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50 55 60

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Ile Ser Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser
115 120

<210> 56
<211> 5
<212> PRT
<213> artificial

<220>
<223> CD33 HCDR1 of AH11
<400> 56

Asn Tyr Gly Met Asn
1 5

<210> 57
<211> 17
<212> PRT
<213> artificial

<220>
<223> CD33 HCDR2 of AH11
<400> 57

Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe Lys
1 5 10 15

Gly

<210> 58
<211> 13
<212> PRT
<213> artificial

<220>
<223> HCDR3
<400> 58

Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr

1

5

10

<210> 59

<211> 366

<212> DNA

<213> artificial

<220>

<223> CD33 VH of AH11

<400> 59

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tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct 120

ccaggacagg gtttaaagtg gatgggctgg ataaacacct acactggaga gccaacatat 180

gctgatgact tcaagggacg ggttaccatg acttcggata cctctaccag cactgcctat 240

atggaaatca gcagcctcag aagtgatgac acggctgtat attactgtgc gcgctggagt 300

tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360

tcctca 366

<210> 60

<211> 113

<212> PRT

<213> artificial

<220>

<223> CD33 VL of AH11

<400> 60

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu Thr Val Ser Leu Gly
1 5 10 15Glu Arg Thr Thr Ile Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser
20 25 30Ser Lys Asn Lys Asn Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln
35 40 45Pro Pro Lys Leu Leu Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile
50 55 60Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr
65 70 75 80Ile Asp Ser Leu Gln Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln
85 90 95Ser Ala His Phe Pro Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile
100 105 110

eof-seq1.txt

Lys

<210> 61
<211> 17
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR1 of AH11

<400> 61

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
1 5 10 15

Ala

<210> 62
<211> 7
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR2 of AH11

<400> 62

Trp Ala Ser Thr Arg Glu Ser
1 5

<210> 63
<211> 9
<212> PRT
<213> artificial

<220>
<223> CD33LCDR3 of AH11

<400> 63

Gln Gln Ser Ala His Phe Pro Ile Thr
1 5

<210> 64
<211> 339
<212> DNA
<213> artificial

<220>
<223> CD33 VL of AH11

<400> 64
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eof-seq1.txt

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tggtaccagc agaaaccagg acagcctcct aaattactcc tttcctgggc atctacgcgg 180
gaatccggga tccctgaccg attcagtggc agcgggtctg ggacagattt cactctcact 240
attgacagcc tgcagcctga agattctgca acttactatt gtcaacagtc tgcccacttc 300
ccgatcacct ttggccaagg gacacgactg gagattaaa 339
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<210> 65

<211> 250

<212> PRT

<213> artificial

<220>

<223> CD33 HL of AH11

<400> 65

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
50 55 60

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Ile Ser Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

eo1f-seq1.txt

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys
245 250

<210> 66
<211> 750
<212> DNA
<213> artificial

<220>
<223> CD33 HL of AH11

<400> 66
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tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct 120
ccaggacagg gtttaaagtg gatgggctgg ataaacacct acactggaga gccaacatat 180
gctgatgact tcaagggacg ggttaccatg acttcggata cctctaccag cactgcctat 240
atggaaaatca gcagcctcag aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcagggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa 750

<210> 67
<211> 505
<212> PRT
<213> artificial

eo1f-seq1.txt

<220>

<223> CD33 AH11 HL x H2C HL

<400> 67

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
50 55 60

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Ile Ser Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

eo1f-seq1.txt

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
 245 250 255
 Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 260 265 270
 Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
 275 280 285
 Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 290 295 300
 Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 305 310 315 320
 Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 325 330 335
 Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 340 345 350
 Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
 355 360 365
 Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 370 375 380
 Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
 385 390 395 400
 Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 405 410 415
 Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
 420 425 430
 Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
 435 440 445
 Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
 450 455 460
 Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
 465 470 475 480
 Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 68
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AH11 HL x H2C HL

<400> 68
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ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
gctgatgact tcaagggacg ggttaccatg acttcggata cctctaccag cactgcctat 240
atggaaatca gcagcctcag aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgctc 360
tcctcagggtg gtggtgggtc tggcggcggc ggctccgggtg gtggtgggtc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactctac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg 780
gtcgagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc 840
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ttggaatggg ttgctcgcac aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaaag acagggtcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
ggtaatagct acatatccta ctgggcttac tggggccaag ggactctggt caccgtctcc 1140
tcagggtgtg gtggttctgg cggcggcggc tccgggtgtg gtggttctca gactgttgtg 1200
actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtggctcc 1260
tcgactgggg ctgttacatc tggctactac ccaaactggg tccaacaaaa accaggtcag 1320
gcaccccggtg gtctaataagg tgggactaag ttctcgccc ccggtactcc tgccagattc 1380
tcaggctccc tgcttggagg caaggctgcc ctaccctct caggggtaca gccagaggat 1440

eof-seq1.txt

gaggcagaat attactgtgc tctatggtac agcaaccgct ggggtgttcgg tggaggaacc 1500
aaactgactg tccta 1515

<210> 69
<211> 505
<212> PRT
<213> artificial

<220>
<223> CD33 AH11 HL x F12Q HL

<400> 69

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
50 55 60

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Ile Ser Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

eo1f-seq1.txt

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

eof-seq1.txt

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 70
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AH11 HL x F12Q HL

<400> 70
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tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct 120
ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
gctgatgact tcaagggacg ggttaccatg acttcggata cctctaccag cactgcctat 240
atggaaaatca gcagcctcag aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcaggtg gtgggtggtc tggcggcggc ggctccggtg gtgggtggtc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggag agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactctac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttgccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg 780
gtcgagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc 840
tctggattca ccttcaatag ctacgccatg aactgggtcc gccaggctcc aggaaagggt 900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaaaag gcagggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080

eolf-seq1.txt

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actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtggctcc 1260
tcgactgggg ctgttacatc tggcaactac ccaaactggg tccaacaaaa accagggtcag 1320
gcaccccggtg gtctaataagg tgggactaag ttcctcgccc ccggtactcc tgccagattc 1380
tcaggctccc tgcttggagg caaggctgcc ctcaccctct caggggtaca gccagaggat 1440
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aaactgactg tccta 1515

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<210> 71
<211> 505
<212> PRT
<213> artificial

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<220>
<223> CD33 AH11 HL x I2C HL

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<400> 71

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Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1          5          10          15

```

```

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20          25          30

```

```

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met
35          40          45

```

```

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe
50          55          60

```

```

Lys Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65          70          75          80

```

```

Met Glu Ile Ser Ser Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85          90          95

```

```

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100          105          110

```

```

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115          120          125

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```

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130          135          140

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eo1f-seq1.txt

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

eof-seq1.txt

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 72
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 AH11 HL x I2C HL

<400> 72
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ccaggacagg gtttaaagtg gatgggctgg ataaacacct aactggaga gccaacatat 180
gctgatgact tcaagggacg gggtaccatg acttcggata cctctaccag cactgcctat 240
atggaaatca gcagcctcag aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
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atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc aagaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactcac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg 780

eof-seq1.txt

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ttggaatggg ttgctcgcac aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaaag acaggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
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tcaggctccc tgcttggagg caaggctgcc ctaccctct caggggtaca gccagaggat 1440
gaggcagaat attactgtgt tctatggtac agcaaccgct ggggtgttcgg tggaggaacc 1500
aaactgactg tccta 1515

<210> 73
<211> 122
<212> PRT
<213> artificial

<220>
<223> CD33 VH of B3

<400> 73

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
50 55 60

Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
Page 60

100

105

110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser
 115 120

<210> 74
 <211> 5
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HCDR1 of B3

<400> 74

Asn Tyr Gly Met Asn
 1 5

<210> 75
 <211> 17
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HCDR2 of B3

<400> 75

Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe Gln
 1 5 10 15

Gly

<210> 76
 <211> 13
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HCDR3 of B3

<400> 76

Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr
 1 5 10

<210> 77
 <211> 366
 <212> DNA
 <213> artificial

<220>
 <223> CD33 VH of B3

<400> 77

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60

eof-seq1.txt

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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gacaaactat 180
gctgataagt tccagggacg cgttaccttc acttcggata cctctaccag cactgcctat 240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctca 366

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<210> 78
 <211> 113
 <212> PRT
 <213> artificial

<220>
 <223> CD33 VL of B3

<400> 78

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Met Thr Val Ser Leu Gly
 1 5 10 15

Glu Arg Thr Thr Ile Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser
 20 25 30

Ser Thr Asn Lys Asn Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln
 35 40 45

Pro Pro Lys Leu Leu Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile
 50 55 60

Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr
 65 70 75 80

Ile Asp Ser Leu Gln Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln
 85 90 95

Ser Ala His Phe Pro Ile Thr Phe Gly Gln Gly Thr Arg Leu Asp Ile
 100 105 110

Lys

<210> 79
 <211> 17
 <212> PRT
 <213> artificial

<220>
 <223> CD33 LCDR1 of B3

eof-seq1.txt

<400> 79

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
1 5 10 15

Ala

<210> 80

<211> 6

<212> PRT

<213> artificial

<220>

<223> CD33 LCDR2 of B3

<400> 80

Trp Ala Ser Thr Arg Glu
1 5

<210> 81

<211> 9

<212> PRT

<213> artificial

<220>

<223> CD33LCDR3 of B3

<400> 81

Gln Gln Ser Ala His Phe Pro Ile Thr
1 5

<210> 82

<211> 339

<212> DNA

<213> artificial

<220>

<223> CD33 VL of B3

<400> 82

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atcaactgca agtccagcca gagtgtttta gacagctcca cgaataagaa ctcttagct	120
tggtaccagc agaaaccagg acagcctcct aaattactcc tttcctgggc atctacgcgg	180
gaatccggga tccctgaccg attcagtggc agcgggtctg ggacagattt cactctcact	240
attgacagcc tgcagcctga agattctgca acttactatt gtcaacagtc tgcccacttc	300
ccgatcacct ttggccaagg gacacgactg gacattaa	339

<210> 83

<211> 250

eof-seq1.txt

<212> PRT
<213> artificial

<220>
<223> CD33 HL of B3

<400> 83

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
50 55 60

Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Met Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

eo1f-seq1.txt

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Asp Ile Lys
245 250

<210> 84

<211> 750

<212> DNA

<213> artificial

<220>

<223> CD33 HL of B3

<400> 84

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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gacaaactat	180
gctgataagt tccagggacg cgttaccttc acttcggata cctctaccag cactgcctat	240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt	300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc	360
tcctcaggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg	420
atgacacagt ctccagactc catgactgtg tctctgggcg agaggaccac catcaactgc	480
aagtcagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag	540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg	600
atcccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc	660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc	720
tttggccaag ggacacgact ggacattaaa	750

<210> 85

<211> 505

<212> PRT

<213> artificial

<220>

<223> CD33 B3 HL x H2C HL

<400> 85

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

eo1f-seq1.txt

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
50 55 60

Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Met Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Asp Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

eo1f-seq1.txt

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 86
<211> 1515
<212> DNA
<213> artificial

<220>

<223> CD33 B3 HL x H2C HL

<400> 86

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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gacaaactat      180
gctgataagt tccagggacg cgttaccttc acttcggata cctctaccag cactgcctat      240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt      300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc      360
tcctcagggtg gtggtgggttc tggcggcggc ggctccgggtg gtggtgggttc tgacatcgtg      420
atgacacagt ctccagactc catgactgtg tctctgggcg agaggaccac catcaactgc      480
aagtccagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag      540
cagaaaccag gacagcctcc taaattactc ctttctctggg catctacgcg ggaatccggg      600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactctac tattgacagc      660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc      720
tttggccaaag ggacacgact ggacattaaa tccggagggtg gtgggtccga ggtgcagctg      780
gtcgagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaaactctc atgtgcagcc      840
tctggattca ccttcaataa gtacgccatg aactgggtcc gccaggctcc aggaaagggt      900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat      960
tcagtgaaaag acagggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa     1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc     1080
ggtaatagct acatatccta ctgggcttac tggggccaag ggactctggt caccgtctcc     1140
tcagggtggtg gtggttcttg cggcggcggc tccggtggtg gtggttctca gactgttgtg     1200
actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtggctcc     1260
tcgactgggg ctgttacatc tggctactac ccaaactggg tccaacaaaa accaggtcag     1320
gcaccccgctg gtctaatagg tgggactaag ttctcgcgcc ccggtactcc tgccagattc     1380
tcaggctccc tgcttgagg caaggctgcc ctaccctct caggggtaca gccagaggat     1440
gaggcagaat attactgtgc tctatggtac agcaaccgct ggggtgttcgg tggaggaacc     1500
aaactgactg tccta                                     1515

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<210> 87

<211> 505

<212> PRT

<213> artificial

<220>

<223> CD33 B3 HL x F12Q HL

eo1f-seq1.txt

<400> 87

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Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20     25     30
Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35     40     45
Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
50     55     60
Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65     70     75     80
Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85     90     95
Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100    105    110
Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115    120    125
Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130    135    140
Pro Asp Ser Met Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145    150    155    160
Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
165    170    175
Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180    185    190
Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195    200    205
Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210    215    220
Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225    230    235    240

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eo1f-seq1.txt

Phe Gly Gln Gly Thr Arg Leu Asp Ile Lys Ser Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

eof-seq1.txt

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 88
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 B3 HL x F12Q HL

<400> 88
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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gacaaactat 180
gctgataagt tccagggacg cggtaccttc acttcggata cctctaccag cactgcctat 240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcagggtg gtggtgggtc tggcggcggc ggctccgggtg gtggtgggtc tgacatcgtg 420
atgacacagt ctccagactc catgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactctac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggacattaaa tccggagggtg gtggctccga ggtgcagctg 780
gtcagagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc 840
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tcagtgaag gcaggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
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tcaggctccc tgcttggagg caaggctgcc ctcaccctct caggggtaca gccagaggat 1440
gaggcagaat attactgtgt tctatggtac agcaaccgct ggggtgttcgg tggaggaacc 1500

aaactgactg tccta

1515

<210> 89

<211> 505

<212> PRT

<213> artificial

<220>

<223> CD33 B3 HL x I2C HL

<400> 89

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
 50 55 60

Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80

Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
 130 135 140

Pro Asp Ser Met Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
 145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
 165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
 180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser


```

195                                200                                205
Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210                                215                                220
Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225                                230                                235                                240
Phe Gly Gln Gly Thr Arg Leu Asp Ile Lys Ser Gly Gly Gly Gly Ser
245                                250                                255
Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260                                265                                270
Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275                                280                                285
Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290                                295                                300
Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305                                310                                315                                320
Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325                                330                                335
Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340                                345                                350
Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355                                360                                365
Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370                                375                                380
Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385                                390                                395                                400
Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405                                410                                415
Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420                                425                                430
Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435                                440                                445

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eof-seq1.txt

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 90
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 B3 HL x I2C HL

<400> 90
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ccaggacagg gtttagagtg gatgggctgg ataaacacct acactggaga gacaaactat 180
gctgataagt tccagggacg cgttaccttc acttcggata cctctaccag cactgcctat 240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcagggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg 420
atgacacagt ctccagactc catgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactctac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggacattaaa tccggagggtg gtggctccga ggtgcagctg 780
gtcgagtctg gaggaggatt ggtgcagcct ggagggcat tgaaactctc atgtgcagcc 840
tctggattca ccttcaataa gtacgccatg aactgggtcc gccaggctcc aggaaaggggt 900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaag acaggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
ggtaatagct acatatccta ctgggcttac tggggccaag ggactctggt caccgtctcc 1140

eof-seq1.txt

tcagggtggtg gtggttctgg cggcggcggc tccggtggtg gtggttctca gactgttgtg 1200
 actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtggctcc 1260
 tcgactgggg ctgttacatc tggcaactac ccaaactggg tccaacaaaa accaggtcag 1320
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 tcaggctccc tgcttggagg caaggctgcc ctcacctct caggggtaca gccagaggat 1440
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 aaactgactg tccta 1515

<210> 91
 <211> 122
 <212> PRT
 <213> artificial

<220>
 <223> CD33 VH of F2

<400> 91

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
 50 55 60

Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80

Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser
 115 120

<210> 92
 <211> 5
 <212> PRT
 <213> artificial

<220>

<223> CD33 HCDR1 of F2

<400> 92

Asn Tyr Gly Met Asn
1 5

<210> 93

<211> 17

<212> PRT

<213> artificial

<220>

<223> CD33 HCDR2 of F2

<400> 93

Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe Gln
1 5 10 15

Gly

<210> 94

<211> 13

<212> PRT

<213> artificial

<220>

<223> CD33 HCDR3 of F2

<400> 94

Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr
1 5 10

<210> 95

<211> 366

<212> DNA

<213> artificial

<220>

<223> CD33 VH of F2

<400> 95

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tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct	120
ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gacaaactat	180
gctgataagt tccaggggacg cgttaccttc acttcggata cctctaccag cactgcctat	240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt	300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc	360
tcctca	366

eo1f-seq1.txt

<210> 96
 <211> 113
 <212> PRT
 <213> artificial

<220>
 <223> CD33 VL of F2

<400> 96

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu Ser Val Ser Leu Gly
 1 5 10 15

Glu Arg Thr Thr Ile Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser
 20 25 30

Ser Thr Asn Lys Asn Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln
 35 40 45

Pro Pro Lys Leu Leu Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile
 50 55 60

Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr
 65 70 75 80

Ile Asp Ser Leu Gln Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln
 85 90 95

Ser Ala His Phe Pro Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile
 100 105 110

Lys

<210> 97
 <211> 17
 <212> PRT
 <213> artificial

<220>
 <223> CD33 LCDR1 of F2

<400> 97

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
 1 5 10 15

Ala

<210> 98
 <211> 7

eof-seq1.txt

<212> PRT
<213> artificial

<220>
<223> CD33 LCDR2 of F2

<400> 98

Trp Ala Ser Thr Arg Glu Ser
1 5

<210> 99
<211> 9
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR3 of F2

<400> 99

Gln Gln Ser Ala His Phe Pro Ile Thr
1 5

<210> 100
<211> 339
<212> DNA
<213> artificial

<220>
<223> CD33 VL of F2

<400> 100
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atcaactgca agtccagcca gagtgtttta gacagctcca cgaataagaa ctcttagct 120
tggtaccagc agaaaccagg acagcctcct aaattactcc tttcctgggc atctacgcgg 180
gaatccggga tccctgaccg attcagtggc agcgggtctg ggacagattt cactctcact 240
attgacagcc tgcagcctga agattctgca acttactatt gtcaacagtc tgcccacttc 300
ccgatcacct ttggccaagg gacacgactg gagattaaa 339

<210> 101
<211> 250
<212> PRT
<213> artificial

<220>
<223> CD33 HL of F2

<400> 101

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr

20

25

30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
 50 55 60

Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80

Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
 130 135 140

Pro Asp Ser Leu Ser Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
 145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
 165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
 180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
 195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
 210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
 225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys
 245 250

<210> 102
 <211> 750
 <212> DNA
 <213> artificial

eof-seq1.txt

<220>

<223> CD33 HL of F2

<400> 102

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tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct    120
ccaggacagg gtttagagtg gatgggctgg ataaacacct acactggaga gacaaactat    180
gctgataagt tccaggggacg cgttaccttc acttcggata cctctaccag cactgcctat    240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt    300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc    360
tcctcagggtg gtgggtggttc tggcggcggc ggctccggtg gtgggtggttc tgacatcgtg    420
atgacacagt ctccagactc cctgtctgtg tctctgggcg agaggaccac catcaactgc    480
aagtcagccc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag    540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg    600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc    660
ctgcagcctg aagattctgc aacttactat tgtaacagct ctgcccactt cccgatcacc    720
tttggccaag ggacacgact ggagattaaa    750

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<210> 103

<211> 505

<212> PRT

<213> artificial

<220>

<223> CD33 F2 HL x H2C HL

<400> 103

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Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1          5          10          15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
          20          25          30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
          35          40          45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
          50          55          60

Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65          70          75          80

Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys

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Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Ser Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

eo1f-seq1.txt

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 104
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 F2 HL x H2C HL

<400> 104
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tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct 120
ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gacaaactat 180
gctgataagt tccagggacg cgttaccttc acttcggata cctctaccag cactgcctat 240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt 300

eof-seq1.txt

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tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcaggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg 420
atgacacagt ctccagactc cctgtctgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcttggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa tccggaggtg gtggctccga ggtgcagctg 780
gtcgagtctg gaggaggatt ggtgcagcct ggagggcat tgaaactctc atgtgcagcc 840
tctggattca ccttcaataa gtacgccatg aactgggtcc gccagggtcc aggaaaggg 900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaag acaggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
ggtaatagct acatatccta ctgggcttac tggggccaag ggactctggt caccgtctcc 1140
tcaggtggtg gtggttctgg cggcggcggc tccggtggtg gtggttctca gactgttggt 1200
actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtggctcc 1260
tcgactgggg ctgttacatc tggctactac ccaaactggg tccaacaaaa accaggtcag 1320
gcaccccggt gtctaatagg tgggactaag ttctctgccc ccggtactcc tgccagattc 1380
tcaggctccc tgcttggagg caaggctgcc ctcaccctct caggggtaca gccagaggat 1440
gaggcagaat attactgtgc tctatggtac agcaaccgct ggggtgttcgg tggaggaacc 1500
aaactgactg tccta 1515

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<210> 105
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 F2 HL x F12Q HL
 <400> 105

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45

eo1f-seq1.txt

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
50 55 60

Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Ser Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val

eo1f-seq1.txt
300

290

295

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 106
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 F2 HL x F12Q HL

eof-seq1.txt

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<400> 106
caggtgcagc tgggtgcagtc tggagctgag gtgaagaagc ctggagagtc agtcaaggct 60
tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct 120
ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gacaaactat 180
gctgataagt tccagggacg cgttaccttc acttcggata cctctaccag cactgcctat 240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcagggtg gtgggtggtc tggcggcggc ggctccgggtg gtgggtggtc tgacatcgtg 420
atgacacagt ctccagactc cctgtctgtg tctctgggcy agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccccgacc gattcagtgg cagcgggtct gggacagatt tcaactctac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg 780
gtcaggtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc 840
tctggattca cttcaatag ctacgccatg aactgggtcc gccaggctcc aggaaaagggt 900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaag gcaggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
ggtaatagct acgtttcctg gtgggcttac tggggccaag ggactctggt caccgtctcc 1140
tcaggtggtg gtggttctgg cggcggcggc tccggtggtg gtggttctca gactgttggt 1200
actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtggctcc 1260
tcgactgggg ctgttacatc tggcaactac ccaaactggg tccaacaaaa accaggtcag 1320
gcaccccggtg gtctaatagg tgggactaag ttcctcgccc ccggtactcc tgccagattc 1380
tcaggctccc tgcttgagg caaggctgcc ctcaccctct caggggtaca gccagaggat 1440
gaggcagaat attactgtgt tctatggtac agcaaccgct ggggtgttcgg tggaggaacc 1500
aaactgactg tccta 1515

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<210> 107
<211> 505
<212> PRT
<213> artificial

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<220>
<223> CD33 F2 HL x I2C HL

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<400> 107

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eo1f-seq1.txt

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15
 Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30
 Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45
 Gly Trp Ile Asn Thr Tyr Thr Gly Glu Thr Asn Tyr Ala Asp Lys Phe
 50 55 60
 Gln Gly Arg Val Thr Phe Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80
 Met Glu Leu Arg Asn Leu Lys Ser Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95
 Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110
 Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125
 Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
 130 135 140
 Pro Asp Ser Leu Ser Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
 145 150 155 160
 Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
 165 170 175
 Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
 180 185 190
 Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
 195 200 205
 Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu
 210 215 220
 Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
 225 230 235 240
 Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
 245 250 255

eo1f-seq1.txt

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 260 265 270
 Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
 275 280 285
 Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 290 295 300
 Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 305 310 315 320
 Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 325 330 335
 Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 340 345 350
 Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
 355 360 365
 Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 370 375 380
 Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
 385 390 395 400
 Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 405 410 415
 Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
 420 425 430
 Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
 435 440 445
 Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
 450 455 460
 Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
 465 470 475 480
 Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
 485 490 495
 Gly Gly Gly Thr Lys Leu Thr Val Leu

<210> 108
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 F2 HL x I2C HL

<400> 108
caggtgcagc tgggtgcagtc tggagctgag gtgaagaagc ctggagagtc agtcaaggct 60
tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct 120
ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gacaaactat 180
gctgataagt tccagggacg cgttaccttc acttcggata cctctaccag cactgcctat 240
atggaactcc gcaacctcaa aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc 360
tcctcagggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg 420
atgacacagt ctccagactc cctgtctgtg tctctgggag agaggaccac catcaactgc 480
aagtcacagg agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgag ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactctac tattgacagc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatacacc 720
tttggccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg 780
gtcgagtctg gaggaggatt ggtgcagcct ggagggtcat tgaaaactctc atgtgcagcc 840
tctggattca ccttcaataa gtacgccatg aactgggtcc gccagggtcc aggaaagggt 900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaag acagggtcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
ggtaatagct acatatccta ctgggcttac tggggccaag ggactctggt caccgtctcc 1140
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gcaccccggtg gtctaataag tgggactaag ttcctcgccc ccggtactcc tgccagattc 1380
tcaggctccc tgcttgagg caaggctgcc ctacacctct caggggtaca gccagaggat 1440
gaggcagaat attactgtgt tctatggtac agcaaccgct ggggtgttcgg tggaggaacc 1500
aaactgactg tccta 1515

eo1f-seq1.txt

<210> 109
 <211> 122
 <212> PRT
 <213> artificial

<220>
 <223> CD33 VH of B10

<400> 109

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
 50 55 60

Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80

Met Glu Ile Arg Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser
 115 120

<210> 110
 <211> 5
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HCDR1 of B10

<400> 110

Asn Tyr Gly Met Asn
 1 5

<210> 111
 <211> 17
 <212> PRT
 <213> artificial

eo1f-seq1.txt

<220>

<223> CD33HCDR2 of B10

<400> 111

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Gly

<210> 112

<211> 13

<212> PRT

<213> artificial

<220>

<223> CD33 HCDR3 of B10

<400> 112

Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr
1 5 10

<210> 113

<211> 366

<212> DNA

<213> artificial

<220>

<223> CD33 VH of B10

<400> 113

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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacctat	180
gctgataagt tccagggacg cgttaccatg actacggata cctctaccag cactgcctat	240
atggaaaatcc gcaacctcag aagtgatgac acggctgtat attactgtgc gcgctggagt	300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgtc	360
tcctca	366

<210> 114

<211> 113

<212> PRT

<213> artificial

<220>

<223> CD33 VL of B10

<400> 114

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu Thr Val Ser Leu Gly
1 5 10 15

eo1f-seq1.txt

Glu Arg Thr Thr Ile Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser
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Ser Asn Asn Lys Asn Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln
35 40 45
Pro Pro Lys Leu Leu Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile
50 55 60
Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr
65 70 75 80
Ile Asp Gly Leu Gln Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln
85 90 95
Ser Ala His Phe Pro Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile
100 105 110

Lys

<210> 115
<211> 17
<212> PRT
<213> artificial
<220>
<223> CD33 LCDR1 of B10
<400> 115

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Asn Asn Lys Asn Ser Leu
1 5 10 15

Ala

<210> 116
<211> 7
<212> PRT
<213> artificial
<220>
<223> CD33 LCDR2 of B10
<400> 116

Trp Ala Ser Thr Arg Glu Ser
1 5

<210> 117

eof-seq1.txt

<211> 9
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<220>
<223> CD33 LCDR3 of B10

<400> 117

Gln Gln Ser Ala His Phe Pro Ile Thr
1 5

<210> 118
<211> 339
<212> DNA
<213> artificial

<220>
<223> CD33 VL of B10

<400> 118
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atcaactgca agtccagcca gagtgtttta gacagctcca acaataagaa ctccttagct 120
tggtaccagc agaaaccagg acagcctcct aaattactcc tttcctgggc atctacgcgg 180
gaatccggga tccctgaccg attcagtggc agcggttctg ggacagattt cactctcact 240
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<210> 119
<211> 250
<212> PRT
<213> artificial

<220>
<223> CD33 HL of B10

<400> 119

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
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Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
50 55 60

Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

eo1f-seq1.txt

Met Glu Ile Arg Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90
Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110
Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125
Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140
Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160
Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Asn Asn Lys Asn Ser Leu
165 170 175
Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190
Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205
Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Gly Leu Gln Pro Glu
210 215 220
Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240
Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys
245 250

<210> 120
<211> 750
<212> DNA
<213> artificial

<220>
<223> CD33 HL of B10

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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacctat 180
gctgataagt tccagggacg cgttaccatg actacggata cctctaccag cactgcctat 240

eof-seq1.txt

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tcctcaggtg gtggtggttc tggcggcggc ggctccgggtg gtggtggttc tgacatcgtg 420
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aagtcagacc agagtgtttt agacagctcc aacaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcggttct gggacagatt tcactctcac tattgacggc 660
ctgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa 750

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<210> 121
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 B10 HL x H2C HL

<400> 121

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
 50 55 60

Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80

Met Glu Ile Arg Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
 130 135 140

eo1f-seq1.txt

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Asn Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Gly Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val

385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

$\langle 210 \rangle$	122
$\langle 211 \rangle$	1515

<215> a l t i t u d i n a l

<220>

<400> 122
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ccaggacagg gtttagactg gatgggctgg ataaacacct acactggaga gccaacct

gctgataaagt tccatgggcaty cgttaccatg attatcggata cctctaccag cactgcct
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ttgagttgaty gttatcatgt ttacttttgac tacttggggcc aaggcactac ggtcaccy
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aagtccagcc agagtgtttt agacagctcc aacaataaga actccttagc ttggtac

tagaaactag gacagctctc taaattactc ctttcttggg catctacgcg ggaatccg
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<210>	122	
<211>	1515	
<212>	DNA	
<213>	artificial	
<220>		
<223>	CD33 B10 HL x H2C HL	
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ccaggacagg	gttttagagt gatgggcttg ataacacct acactggaga gccaacctat	180
gctgataagt	tccagggacg cgttaccatg actacggata cctctaccag cactgcctat	240
atggaaatcc	gcaacctcag aagtgatgac acggctgtat attactgtgc gcgctggagt	300
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tcctcaggtg	gtggtgggtc tggcggcggc ggctccggtg gtggtgggtc tgacatcgtg	420
atgacacagt	ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc	480
aagtccagcc	agagtgtttt agacagctcc aacaataaga actccttagc ttggtaccag	540
cagaaaccag	gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg	600
atccctgacc	gattcagtgg cagcggttct gggacagatt tcactctcac tattgacggc	660
ctgcagcctg	aaqattctgc aaacttactat tgtcaacagt ctgccactt cccgatcacc	720

eof-seq1.txt

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gtcgaagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc 840
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ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaaaag acagggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
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aaactgactg tccta 1515

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<210> 123
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 B10 HL x F12Q HL

<400> 123

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
50 55 60

Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Ile Arg Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

eo1f-seq1.txt

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Asn Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Gly Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

eo1f-seq1.txt

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 124
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 B10 HL x F12Q HL

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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacctat 180
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atggaaaatcc gcaacctcag aagtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcactac ggtcaccgctc 360

eolf-seq1.txt

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aaactgactg tccta	1515

<210> 125
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 B10 HL x I2C HL

<400> 125

Gln Val	Gln Leu Val	Gln Ser Gly Ala	Glu Val	Lys Lys Pro	Gly Glu
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Ser Val	Lys Val	Ser Cys Lys Ala	Ser Gly Tyr Thr	Phe Thr	Asn Tyr
	20	25		30	

Gly Met	Asn Trp Val	Lys Gln Ala	Pro Gly Gln Gly	Leu Glu Trp	Met
	35	40		45	

eo1f-seq1.txt

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
50 55 60

Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Ile Arg Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100 105 110

Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Asn Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Gly Leu Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

eof-seq1.txt

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 126
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 B10 HL x I2C HL

<400> 126
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eof-seq1.txt

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gctgataagt tccagggacg cgttaccatg actacggata cctctaccag cactgcctat 240
atggaaatcc gcaacctcag aagtgatgac acggctgtat attactgtgc gcgctggagt 300
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gcaccccggt gtctaataag tgggactaag ttctcgccc ccggtactcc tgccagattc 1380
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aaactgactg tccta 1515

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<210> 127
 <211> 122
 <212> PRT
 <213> artificial

<220>
 <223> CD33 VH of E11

<400> 127

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu

1 5 10 15
 Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30
 Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45
 Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
 50 55 60
 Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80
 Met Glu Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95
 Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110
 Gly Gln Gly Thr Ser Val Thr Val Ser Ser
 115 120

<210> 128
 <211> 5
 <212> PRT
 <213> artificial

 <220>
 <223> CD33 HCDR1 of E11

 <400> 128

Asn Tyr Gly Met Asn
1 5

<210> 129
 <211> 17
 <212> PRT
 <213> artificial

 <220>
 <223> CD33 HCDR2 of E11

 <400> 129

Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe Gln
1 5 10 15

Gly

eof-seq1.txt

<210> 130
 <211> 13
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HCDR3 of E11

<400> 130

Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr
 1 5 10

<210> 131
 <211> 366
 <212> DNA
 <213> artificial

<220>
 <223> CD33 VH of E11

<400> 131
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 ccaggacagg gtttagagtg gatgggctgg ataaacacct acactggaga gccaacctat 180
 gctgataagt tccaggggacg cgttaccatg actacggata cctctaccag cactgcctat 240
 atggaaatcc gcaacctcgg aggtgatgac acggctgtat attactgtgc gcgctggagt 300
 tggagtgatg gttactacgt ttactttgac tactggggcc aaggcacttc ggtcaccgctc 360
 tcctca 366

<210> 132
 <211> 113
 <212> PRT
 <213> artificial

<220>
 <223> CD33 VL of E11

<400> 132

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu Thr Val Ser Leu Gly
 1 5 10 15

Glu Arg Thr Thr Ile Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser
 20 25 30

Ser Thr Asn Lys Asn Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln
 35 40 45

Pro Pro Lys Leu Leu Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile
 50 55 60

eo1f-seq1.txt

Pro Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr
65 70 75 80

Ile Asp Ser Pro Gln Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln
85 90 95

Ser Ala His Phe Pro Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile
100 105 110

Lys

<210> 133
<211> 17
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR1 of E11

<400> 133

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
1 5 10 15

Ala

<210> 134
<211> 7
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR2 of E11

<400> 134

Trp Ala Ser Thr Arg Glu Ser
1 5

<210> 135
<211> 9
<212> PRT
<213> artificial

<220>
<223> CD33 LCDR3 of E11

<400> 135

Gln Gln Ser Ala His Phe Pro Ile Thr
1 5

eof-seq1.txt

<210> 136
 <211> 339
 <212> DNA
 <213> artificial

<220>
 <223> CD33 VL of E11

<400> 136
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 tggtagcagc agaaaccagg acagcctcct aaattactcc tttcctgggc atctacgcgg 180
 gaatccggga tccctgaccg attcagtggc agcgggtctg ggacagattt cactctcact 240
 attgacagcc cgcagcctga agattctgca acttactatt gtcaacagtc tgcccacttc 300
 ccgatcacct ttggccaagg gacacgactg gagattaaa 339

<210> 137
 <211> 250
 <212> PRT
 <213> artificial

<220>
 <223> CD33 HL of E11

<400> 137
 Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15
 Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30
 Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45
 Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
 50 55 60
 Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80
 Met Glu Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95
 Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110
 Gly Gln Gly Thr Ser Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125

eo1f-seq1.txt

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Pro Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys
245 250

<210> 138

<211> 750

<212> DNA

<213> artificial

<220>

<223> CD33 HL of E11

<400> 138

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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacctat	180
gctgataagt tccagggacg cgttaccatg actacggata cctctaccag cactgcctat	240
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tggagtgatg gttactacgt ttactttgac tactggggcc aaggcacttc ggtcaccgtc	360
tcctcaggtg gtgggtggtc tggcggcggc ggctccggtg gtgggtggtc tgacatcgtg	420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc	480
aagtccagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag	540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg	600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcactctcac tattgacagc	660

eof-seq1.txt

ccgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
 tttggccaag ggacacgact ggagattaaa 750

<210> 139
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 E11 HL x H2C HL

<400> 139

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
 1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
 20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
 50 55 60

Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80

Met Glu Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Ser Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125

Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
 130 135 140

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
 145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
 165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
 180 185 190

eo1f-seq1.txt

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Pro Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

eof-seq1.txt

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 140
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 E11 HL x H2C HL

<400> 140
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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacctat 180
gctgataagt tccagggacg cggtaccatg actacggata cctctaccag cactgcctat 240
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tcctcaggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggag agaggaccac catcaactgc 480
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cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgag ggaatccggg 600
atccctgacc gattcagtggt cagcgggtct gggacagatt tcaactctac tattgacagc 660
ccgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatacc 720
tttgccaag ggacacgact ggagattaaa tccggaggtg gtggctccga ggtgcagctg 780
gtcgagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc 840
tctggattca ccttcaataa gtacgccatg aactgggtcc gccaggctcc aggaaagggt 900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaaag acaggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080

eolf-seq1.txt

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actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtggctcc 1260
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gcaccccggtg gtctaataagg tgggactaag ttcctcgccc ccggtactcc tgccagattc 1380
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aaactgactg tccta 1515

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<210> 141
<211> 505
<212> PRT
<213> artificial

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<220>
<223> CD33 E11 HL x F12Q HL

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<400> 141

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Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1      5      10     15

```

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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20     25     30

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```

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35     40     45

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```

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
50     55     60

```

```

Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
65     70     75     80

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```

Met Glu Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr Tyr Cys
85     90     95

```

```

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
100    105    110

```

```

Gly Gln Gly Thr Ser Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
115    120    125

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Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
130    135    140

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eo1f-seq1.txt

Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
145 150 155 160

Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
165 170 175

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
180 185 190

Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
195 200 205

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Pro Gln Pro Glu
210 215 220

Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
225 230 235 240

Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
245 250 255

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
260 265 270

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
275 280 285

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
290 295 300

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
305 310 315 320

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
325 330 335

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
340 345 350

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

eof-seq1.txt

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 142
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 E11 HL x F12Q HL

<400> 142
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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacctat 180
gctgataagt tccagggacg cgttaccatg actacggata cctctaccag cactgcctat 240
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tggagtgatg gttactacgt ttactttgac tactggggcc aaggcacttc ggtcaccgtc 360
tcctcagggtg gtggtgggttc tggcggcggc ggctccgggtg gtggtgggttc tgacatcgtg 420
atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc 480
aagtccagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag 540
cagaaaccag gacagcctcc taaattactc ctttcctggg catctacgcg ggaatccggg 600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactcac tattgacagc 660
ccgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc 720
tttggccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg 780

eof-seq1.txt

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ttggaatggg ttgctcgcac aagaagtaaa tataataatt atgcaacata ttatgccgat 960
tcagtgaaag gcagggtcac catctccaga gatgattcaa aaaacactgc ctatctacaa 1020
atgaacaact tgaaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc 1080
ggtaatagct acgtttcctg gtgggcttac tggggccaag ggactctggg caccgtctcc 1140
tcagggtggg gtggttctgg cggcggcggc tccgggtggg gtggttctca gactgttggt 1200
actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtgggtcc 1260
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gcaccccggt gtctaataagg tgggactaag ttctcgtccc ccggtactcc tgccagattc 1380
tcagggtccc tgcttggagg caaggctgcc ctaccctct caggggtaca gccagaggat 1440
gaggcagaat attactgtgt tctatgggtac agcaaccgct ggggtgttcgg tggaggaacc 1500
aaactgactg tccta 1515

<210> 143
<211> 505
<212> PRT
<213> artificial

<220>
<223> CD33 E11 HL x I2C HL

<400> 143

Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu
1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr
20 25 30

Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
50 55 60

Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
Page 116

eo1f-seq1.txt

100		105		110
Gly	Gln	Gly	Thr	Ser
	115		Val	Thr
			120	Val
Ser	Ser	Gly	Gly	Gly
			125	Gly
Ser	Gly			
Gly	Gly	Gly	Gly	Ser
130			135	Ser
				Asp
Ile	Val	Met	Thr	Gln
	140			Ser
Pro	Asp	Ser	Leu	Thr
145			Val	Ser
			150	Leu
Gly	Glu	Arg	Thr	Thr
		155		Ile
Asn	Cys			
160				
Lys	Ser	Ser	Gln	Ser
			165	Val
Leu	Asp	Ser		
		170		Ser
Thr	Asn	Lys	Asn	Ser
			175	Leu
Ala	Trp	Tyr	Gln	Gln
			180	Lys
Pro	Gly	Gln	Pro	Pro
		185		Lys
Leu	Leu	Leu	Ser	
		190		
Trp	Ala	Ser	Thr	Arg
		195		Glu
Ser	Gly	Ile	Pro	Asp
		200		Arg
Phe	Ser	Gly	Ser	
		205		
Gly	Ser	Gly	Thr	Asp
		210		Phe
Thr	Asp	Phe	Thr	Leu
		215		Thr
Ile	Asp	Ser	Pro	Gln
		220		Pro
Glu				
Asp	Ser	Ala	Thr	Tyr
225				Tyr
			230	Cys
Gln	Gln	Ser	Ala	His
			235	Phe
Pro	Ile	Thr		
		240		
Phe	Gly	Gln	Gly	Thr
		245		Arg
Leu	Glu	Ile	Lys	Ser
			250	Gly
Gly	Gly	Gly	Gly	Gly
			255	Ser
Glu	Val	Gln	Leu	Val
		260		Glu
Ser	Gly	Gly	Gly	Leu
		265		Val
Gln	Pro	Gly	Gly	
		270		
Ser	Leu	Lys	Leu	Ser
		275		Cys
Ala	Ala	Ser	Gly	Phe
		280		Thr
Phe	Asn	Lys	Tyr	
		285		
Ala	Met	Asn	Trp	Val
		290		Arg
Gln	Ala	Pro	Gly	Lys
		295		Gly
Leu	Glu	Trp	Val	
		300		
Ala	Arg	Ile	Arg	Ser
305				Lys
			310	Tyr
Asn	Asn	Tyr	Ala	Thr
			315	Tyr
Tyr	Ala	Asp		
		320		
Ser	Val	Lys	Asp	Arg
			325	Phe
Thr	Ile	Ser	Arg	Asp
			330	Asp
Ser	Lys	Asn	Thr	
		335		
Ala	Tyr	Leu	Gln	Met
		340		Asn
Asn	Leu	Lys	Thr	Glu
		345		Asp
Thr	Ala	Val	Tyr	
		350		

eo1f-seq1.txt

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
355 360 365

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
370 375 380

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
385 390 395 400

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
405 410 415

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
420 425 430

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
435 440 445

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
450 455 460

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
465 470 475 480

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
485 490 495

Gly Gly Gly Thr Lys Leu Thr Val Leu
500 505

<210> 144
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 E11 HL x I2C HL

<400> 144
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tcctgcaagg ctagcgggta taccttcaca aactatggaa tgaactgggt gaagcaggct 120
ccaggacagg gtttagagtg gatgggctgg ataaacacct acactggaga gccaacctat 180
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atggaaaatcc gcaacctcgg aggtgatgac acggctgtat attactgtgc gcgctggagt 300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcacttc ggtcaccgtc 360
tcctcaggtg gtggtggttc tggcggcggc ggctccggtg gtggtggttc tgacatcgtg 420

eolf-seq1.txt

atgacacagt ctccagactc cctgactgtg tctctgggcg agaggaccac catcaactgc	480
aagtccagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag	540
cagaaaccag gacagcctcc taaattactc ctttctctggg catctacgcg ggaatccggg	600
atccctgacc gattcagtgg cagcgggtct gggacagatt tcaactctcac tattgacagc	660
ccgcagcctg aagattctgc aacttactat tgtcaacagt ctgcccactt cccgatcacc	720
tttggccaag ggacacgact ggagattaaa tccggagggtg gtggctccga ggtgcagctg	780
gtcgagtctg gaggaggatt ggtgcagcct ggaggggtcat tgaaactctc atgtgcagcc	840
tctggattca ccttcaataa gtacgccatg aactgggtcc gccagggtcc aggaaagggt	900
ttggaatggg ttgctcgcat aagaagtaaa tataataatt atgcaacata ttatgccgat	960
tcagtgaaaag acagggttcac catctccaga gatgattcaa aaaacactgc ctatctacaa	1020
atgaacaact tgaaaactga ggacactgcc gtgtactact gtgtgagaca tgggaacttc	1080
ggtaatagct acatatccta ctgggcttac tggggccaag ggactctggt caccgtctcc	1140
tcagggtggtg gtggttctgg cggcggcggc tccggtggtg gtggttctca gactgttggtg	1200
actcaggaac cttcactcac cgtatcacct ggtggaacag tcacactcac ttgtgggtcc	1260
tcgactgggg ctgttacatc tggcaactac ccaaactggg tccaacaaaa accaggtcag	1320
gcaccccggtg gtctaataagg tgggactaag ttctctgccc ccggtactcc tgccagattc	1380
tcaggctccc tgcttgagg caaggctgcc ctcaccctct caggggtaca gccagaggat	1440
gaggcagaat attactgtgt tctatggtac agcaaccgct ggggtgttcgg tggaggaacc	1500
aaactgactg tccta	1515

<210> 145
 <211> 1125
 <212> DNA
 <213> human

<400> 145

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ccaaatttct ggctgcaagt gcaggagtca gtgacggtagc aggagggttt gtgcgtcctc	120
gtgccctgca ctttcttcca tcccatacc tactacgaca agaactcccc agttcatggt	180
tactggttcc gggaaggagc cattatatcc ggggactctc cagtggccac aaacaagcta	240
gatcaagaag tacaggagga gactcagggc agattccgcc tccttgggga tcccagtagg	300
aacaactgct ccctgagcat cgtagacgcc aggaggaggg ataatggttc atacttcttt	360
cggatggaga gaggaagtac caaatacagt taaaaatctc cccagctctc tgtgcatgtg	420
acagacttga cccacaggcc caaaatcctc atccctggca ctctagaacc cggccactcc	480
aaaaacctga cctgctctgt gtcctgggccc tgtgagcagg gaacaccccc gatcttctcc	540

eof-seq1.txt

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tggttgtcag ctgccccac ctccctgggc cccaggacta ctcactcctc ggtgctcata 600
atcacccac ggccccagga ccacggcacc aacctgacct gtcaggtgaa gttcgctgga 660
gctggtgtga ctacggagag aaccatccag ctcaacgtca cctatgttcc acagaacca 720
acaactggta tctttccagg agatggctca gggaaacaag agaccagagc aggagtgggt 780
catggggcca ttggaggagc tgggtgtaca gccctgctcg ctctttgtct ctgcctcatc 840
ttcttcatag tgaagacca caggaggaaa gcagccagga cagcagtggg caggaatgac 900
accacccta ccacagggtc agcctccccg aaacaccaga agaagtccaa gttacatggc 960
cccactgaaa cctcaagctg ttcagggtgcc gccctactg tggagatgga tgaggagctg 1020
cattatgctt cctcaactt tcattgggatg aatccttcca aggacacctc caccgaatac 1080
tcagagggtca ggaccagtc cgggcatcat caccatcatc attga 1125

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<210> 146
 <211> 374
 <212> PRT
 <213> human

<400> 146

Met Gly Trp Ser Cys Ile Ile Leu Phe Leu Val Ala Thr Ala Thr Gly
1 5 10 15

Val His Ser Asp Pro Asn Phe Trp Leu Gln Val Gln Glu Ser Val Thr
20 25 30

Val Gln Glu Gly Leu Cys Val Leu Val Pro Cys Thr Phe Phe His Pro
35 40 45

Ile Pro Tyr Tyr Asp Lys Asn Ser Pro Val His Gly Tyr Trp Phe Arg
50 55 60

Glu Gly Ala Ile Ile Ser Gly Asp Ser Pro Val Ala Thr Asn Lys Leu
65 70 75 80

Asp Gln Glu Val Gln Glu Glu Thr Gln Gly Arg Phe Arg Leu Leu Gly
85 90 95

Asp Pro Ser Arg Asn Asn Cys Ser Leu Ser Ile Val Asp Ala Arg Arg
100 105 110

Arg Asp Asn Gly Ser Tyr Phe Phe Arg Met Glu Arg Gly Ser Thr Lys
115 120 125

Tyr Ser Tyr Lys Ser Pro Gln Leu Ser Val His Val Thr Asp Leu Thr
130 135 140

eo1f-seq1.txt

His Arg Pro Lys Ile Leu Ile Pro Gly Thr Leu Glu Pro Gly His Ser
145 150 155 160

Lys Asn Leu Thr Cys Ser Val Ser Trp Ala Cys Glu Gln Gly Thr Pro
165 170 175

Pro Ile Phe Ser Trp Leu Ser Ala Ala Pro Thr Ser Leu Gly Pro Arg
180 185 190

Thr Thr His Ser Ser Val Leu Ile Ile Thr Pro Arg Pro Gln Asp His
195 200 205

Gly Thr Asn Leu Thr Cys Gln Val Lys Phe Ala Gly Ala Gly Val Thr
210 215 220

Thr Glu Arg Thr Ile Gln Leu Asn Val Thr Tyr Val Pro Gln Asn Pro
225 230 235 240

Thr Thr Gly Ile Phe Pro Gly Asp Gly Ser Gly Lys Gln Glu Thr Arg
245 250 255

Ala Gly Val Val His Gly Ala Ile Gly Gly Ala Gly Val Thr Ala Leu
260 265 270

Leu Ala Leu Cys Leu Cys Leu Ile Phe Phe Ile Val Lys Thr His Arg
275 280 285

Arg Lys Ala Ala Arg Thr Ala Val Gly Arg Asn Asp Thr His Pro Thr
290 295 300

Thr Gly Ser Ala Ser Pro Lys His Gln Lys Lys Ser Lys Leu His Gly
305 310 315 320

Pro Thr Glu Thr Ser Ser Cys Ser Gly Ala Ala Pro Thr Val Glu Met
325 330 335

Asp Glu Glu Leu His Tyr Ala Ser Leu Asn Phe His Gly Met Asn Pro
340 345 350

Ser Lys Asp Thr Ser Thr Glu Tyr Ser Glu Val Arg Thr Gln Ser Gly
355 360 365

His His His His His His
370

<210> 147
<211> 1083

eof-seq1.txt

<212> DNA
<213> macaque

<400> 147
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tgcactttct tccatcccgt accctaccac accaggaatt cccagttca tggttactgg 180
ttccgggaag gagccattgt atccttggac tctccagtgg ccacaaacaa gctagatcaa 240
gaagtacagg aggagacca gggccgattc cgcctccttg gggatcccag taggaacaac 300
tgctccctga gcatcgtaga tgccaggagg agggataacg gttcatactt ctttcggatg 360
gagaaaggaa gtaccaaata cagttacaaa tctaccagc tctctgtgca tgtgacagac 420
ttgaccaca ggccccaaat cctcatccct ggagccctag accctgacca ctccaaaaac 480
ctgacctgct ctgtgccctg ggctgtgag cagggaacac ctccaatctt ctctggatg 540
tcagctgccc ccacctccct gggcctcagg accactcact cctcgggtgct cataatcacc 600
ccacggcccc aggaccacgg caccaacctc acctgtcagg tgaagttccc tggagctggc 660
gtgaccacgg agagaaccat ccagctcaat gtctcctatg cttcacagaa cccaagaact 720
gatattttc taggagacgg ctcagggaac caaggagtgg ttcaggagc catcggggga 780
gctggtgtca cagtcctgct cgctctttgt ctctgcctca tcttcttcac agtgaagact 840
cacaggagga aagcagccag gacagcagtg ggcaggatcg acaccacccc cgccacaggg 900
ccaacatcct cgaaacacca gaagaagtcc aagttacatg gcgccactga aacctcaggc 960
tgttcaggta ccacccttac tgtggagatg gatgaggagc tgcactacgc ttccctcaac 1020
tttcatggga tgaatccttc tgaggacacc tccaccgaat actcagaggt caggacccag 1080
tga 1083

<210> 148
<211> 505
<212> PRT
<213> artificial

<220>
<223> CD33 UD H2C HL x AF5 HL
<400> 148

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15
Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
20 25 30
Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

eo1f-seq1.txt

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu Ser Gly Gly Gly Gly Ser Gln
245 250 255

Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ala Ser
260 265 270

Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr Gly
275 280 285

Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met Gly

eo1f-seq1.txt
300

290

295

Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe Lys
305 310 315 320

Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr Leu
325 330 335

Glu Leu His Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys Ala
340 345 350

Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp Gly
355 360 365

Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly
370 375 380

Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser Pro
385 390 395 400

Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys Lys
405 410 415

Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu Ala
420 425 430

Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser Trp
435 440 445

Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser Gly
450 455 460

Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu Asp
465 470 475 480

Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr Phe
485 490 495

Gly Gln Gly Thr Arg Leu Glu Ile Lys
500 505

<210> 149
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 UD H2C HL x AF5 HL

eof-seq1.txt

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<400> 149
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tcattgtgcag cctctggatt caccttcaat aagtacgcc tgaactgggt ccgccaggct 120
ccaggaaagg gtttggaatg ggttgctcgc ataagaagta aatataataa ttatgcaaca 180
tattatgccg attcagtgaag agacagggtc accatctcca gagatgattc aaaaaaact 240
gcctatctac aaatgaacaa cttgaaaact gaggacactg ccgtgtacta ctgtgtgaga 300
catgggaact tcggtaatag ctacatatcc tactgggctt actggggcca agggactctg 360
gtcaccgtct cctcagggtg tgggtggttct ggcgccggcg gctccgggtg tgggtggttct 420
cagactgttg tgactcagga accttcactc accgtatcac ctggtggaac agtcacactc 480
acttgtggct cctcgactgg ggctgttaca tctggctact acccaaactg ggtccaacaa 540
aaaccagggtc aggcaccccg tgggtctaata ggtgggacta agttcctcgc ccccggtact 600
cctgccagat tctcagggtc cctgcttgga ggcaaggctg ccctcaccct ctcaggggta 660
cagccagagg atgaggcaga atattactgt gctctatggt acagcaaccg ctgggtgttc 720
ggtggaggaa ccaaactgac tgtcctatcc ggagggtgtg gctcccagggt gcagctggtc 780
cagtctggag ctgagggtga gaagcctgga gcgtcagtcagggtctcctg caaggctagc 840
gggtatacct tcacaaacta tggaatgaac tgggtgaagc aggtccagg acagggttta 900
aagtggatgg gctggataaa cacctacact ggagagccaa catatgctga tgacttcaag 960
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tacgtttact ttgactactg gggccaaggc actacggta ccgtctcctc aggtgggtgt 1140
ggttctggcg gcggcggtc cggtgggtgt ggttctgaca tcgtgatgac acagtctcca 1200
gactccctga ctgtgtctct gggcgagagg accaccatca actgcaagtc cagccagagt 1260
gttttagaca gctccaagaa taagaactcc ttagcttgg accagcagaa accaggacag 1320
cctcctaaat tactccttct ctgggcatct acgcgggaat ccgggatccc tgaccgattc 1380
agtggcagcg ggtctgggac agatttcact ctactattg acagcctgca gcctgaagat 1440
tctgcaactt actattgtca acagtctgcc cacttcccga tcacctttgg ccaagggaca 1500
cgactggaga ttaaa 1515

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<210> 150
<211> 505
<212> PRT
<213> artificial

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<220>
<223> CD33 UD F12Q HL x AF5 HL
<400> 150

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eo1f-seq1.txt

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15
 Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
 20 25 30
 Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45
 Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60
 Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80
 Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95
 Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
 100 105 110
 Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 115 120 125
 Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
 130 135 140
 Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 145 150 155 160
 Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
 165 170 175
 Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
 180 185 190
 Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
 195 200 205
 Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
 210 215 220
 Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
 225 230 235 240
 Gly Gly Gly Thr Lys Leu Thr Val Leu Ser Gly Gly Gly Gly Ser Gln
 245 250 255

eo1f-seq1.txt

Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ala Ser
 260 265 270
 Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr Gly
 275 280 285
 Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met Gly
 290 295 300
 Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe Lys
 305 310 315 320
 Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr Leu
 325 330 335
 Glu Leu His Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys Ala
 340 345 350
 Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp Gly
 355 360 365
 Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly
 370 375 380
 Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser Pro
 385 390 395 400
 Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys Lys
 405 410 415
 Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu Ala
 420 425 430
 Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser Trp
 435 440 445
 Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser Gly
 450 455 460
 Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu Asp
 465 470 475 480
 Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr Phe
 485 490 495
 Gly Gln Gly Thr Arg Leu Glu Ile Lys

<210> 151
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 UD F12Q HL x AF5 HL

<400> 151
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tcatgtgcag cctctggatt caccctcaat agctacgcc tgaactgggt ccgccagggt 120
ccaggaaagg gtttggaatg ggttgctcgc ataagaagta aatataataa ttatgcaaca 180
tattatgccg attcagtgaaggagggttc accatctcca gagatgattc aaaaaaact 240
gcctatctac aaatgaacaa cttgaaaact gaggacactg ccgtgtacta ctgtgtgaga 300
catgggaact tcggtaatag ctacgtttcc tgggtgggctt actggggcca agggactctg 360
gtcaccgtct cctcagggtg tgggtggttct ggcgccggcg gctccgggtg tgggtggttct 420
cagactgttg tgactcagga acccttcaact accgtatcac ctggtggaac agtcacactc 480
acttggtggct cctcgactgg ggctgttaca tctggcaact acccaaactg ggtccaacaa 540
aaaccagggtc aggcaccccc tgggtctaata ggtgggacta agttcctcgc ccccggtact 600
cctgccagat tctcaggctc cctgcttgga ggcaaggctg ccctcaccct ctcaggggta 660
cagccagagg atgaggcaga atattactgt gttctatggt acagcaaccg ctgggtgttc 720
ggtggaggaa ccaaactgac tgtcctatcc ggagggtggtg gctcccagggt gcagctggtc 780
cagtctggag ctgaggtgaa gaagcctgga gcgtcagtcagggtctcctg caaggctagc 840
gggtatacct tcacaaacta tgggaatgaac tgggtgaagc aggtccagg acaggggtta 900
aagtggatgg gctggataaa cacctacact ggagagccaa catatgctga tgacttcaag 960
ggacgggtta ccatgacttc ggatacctct accagcactg cctatttgga actccacaac 1020
ctcagaagt atgacacggc tgtatattac tgtgcgcgct ggagttggag tgatggttac 1080
tacgtttact ttgactactg gggccaaggc actacggtc cgtctcctc aggtggtggt 1140
ggttctggcg gcggcggctc cgggtggtggt ggttctgaca tcgtgatgac acagtctcca 1200
gactccctga ctgtgtctct gggcgagagg accaccatca actgcaagtc cagccagagt 1260
gttttagaca gctccaagaa taagaactcc ttagcttgggt accagcagaa accaggacag 1320
cctcctaaat tactcctttc ctgggcatct acgcgggaat ccgggatccc tgaccgattc 1380
agtggcagcg ggtctgggac agatttcaact ctactattg acagcctgca gcctgaagat 1440
tctgcaactt actattgtca acagtctgcc cacttcccga tcacctttgg ccaagggaca 1500
cgactggaga ttaaa 1515

eo1f-seq1.txt

<210> 152
 <211> 505
 <212> PRT
 <213> artificial

<220>
 <223> CD33 UD I2C HL x AF5 HL

<400> 152

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
 100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
 130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
 165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
 180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
 195 200 205

eo1f-seq1.txt

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
 210 215 220
 Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
 225 230 235 240
 Gly Gly Gly Thr Lys Leu Thr Val Leu Ser Gly Gly Gly Ser Gln
 245 250 255
 Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ala Ser
 260 265 270
 Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr Gly
 275 280 285
 Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Lys Trp Met Gly
 290 295 300
 Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Asp Phe Lys
 305 310 315 320
 Gly Arg Val Thr Met Thr Ser Asp Thr Ser Thr Ser Thr Ala Tyr Leu
 325 330 335
 Glu Leu His Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Tyr Cys Ala
 340 345 350
 Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp Gly
 355 360 365
 Gln Gly Thr Thr Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly
 370 375 380
 Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser Pro
 385 390 395 400
 Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys Lys
 405 410 415
 Ser Ser Gln Ser Val Leu Asp Ser Ser Lys Asn Lys Asn Ser Leu Ala
 420 425 430
 Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser Trp
 435 440 445
 Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser Gly
 450 455 460

eof-seq1.txt

Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Leu Gln Pro Glu Asp
465 470 475 480

Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr Phe
485 490 495

Gly Gln Gly Thr Arg Leu Glu Ile Lys
500 505

<210> 153
<211> 1515
<212> DNA
<213> artificial

<220>
<223> CD33 UD I2C HL x AF5 HL

<400> 153
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tcatgtgcag cctctggatt caccttcaat aagtacgcca tgaactgggt ccgccaggct 120
ccaggaaagg gtttggaatg ggttgctcgc ataagaagta aatataataa ttatgcaaca 180
tattatgccg attcagtga agacagggtc accatctcca gagatgattc aaaaaacact 240
gcctatctac aaatgaacaa cttgaaaact gaggacactg ccgtgtacta ctgtgtgaga 300
catgggaact tcggtaatag ctacatatcc tactgggctt actggggcca agggactctg 360
gtcaccgtct cctcagggtg tggtggttct ggcggcggcg gctccgggtg tggtggttct 420
cagactgttg tgactcagga accttcaact accgtatcac ctggtggaac agtcacactc 480
acttgtggct cctcgactgg ggctgttaca tctggcaact acccaaactg ggtccaacaa 540
aaaccagggtc aggcaccccc tgggtctaata ggtgggacta agttcctcgc ccccggtact 600
cctgccagat tctcaggctc cctgcttga ggcaaggctg ccctcaccct ctcaagggtgta 660
cagccagagg atgaggcaga atattactgt gttctatggt acagcaaccg ctgggtgttc 720
ggtggaggaa ccaaactgac tgtcctatcc ggagggtggt gctcccagggt gcagctgggtc 780
cagtctggag ctgagggtga gaagcctgga gcgtcagtca aggtctcctg caaggctagc 840
gggtatacct tcacaaacta tggaatgaac tgggtgaagc aggtctcagg acaggggtta 900
aagtggatgg gctggataaa cacctacact ggagagccaa catatgctga tgacttcaag 960
ggacgggtta ccatgacttc ggatacctct accagcactg cctatttga actccacaac 1020
ctcagaagt atgacacggc tgtatattac tgtgcgcgct ggagttggag tgatgggttac 1080
tacgtttact ttgactactg gggccaaggc actacgggtca ccgtctcctc aggtggtggt 1140
ggttctggcg gcggcggctc cggtggtggt ggttctgaca tcgtgatgac acagtctcca 1200

eof-seq1.txt

gactccctga ctgtgtctct gggcgagagg accaccatca actgcaagtc cagccagagt 1260
 gttttagaca gctccaagaa taagaactcc ttagcttggg accagcagaa accaggacag 1320
 cctcctaaat tactcctttc ctgggcatct acgcgggaat ccgggatccc tgaccgattc 1380
 agtggcagcg ggtctgggac agatttcact ctactattg acagcctgca gcctgaagat 1440
 tctgcaactt actattgtca acagtctgcc cacttcccga tcacctttgg ccaagggaca 1500
 cgactggaga ttaaa 1515

<210> 154
 <211> 109
 <212> PRT
 <213> artificial

<220>
 <223> VL of F6A

<400> 154

Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
 1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
 20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
 35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
 50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
 65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
 85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
 100 105

<210> 155
 <211> 125
 <212> PRT
 <213> artificial

<220>
 <223> VH of F6A

<400> 155

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

eo1f-seq1.txt

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ile Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Ser Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Phe Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 156
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL of F6A

<400> 156

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ile Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Ser Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

eo1f-seq1.txt

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Phe Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 157
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL-P of F6A

<400> 157

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

eo1f-seq1.txt

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 158

<211> 125

<212> PRT

<213> artificial

<220>

<223> VH-P of F6A

<400> 158

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ile Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Ser Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Phe Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 159

<211> 249

<212> PRT

<213> artificial

eo1f-seq1.txt

<220>

<223> VH-VL-P of F6A

<400> 159

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ile Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Ser Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Phe Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

eo1f-seq1.txt

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 160
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL of H2C

<400> 160

Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 161
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH of H2C

<400> 161

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val

35

40

45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
 100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
 115 120 125

<210> 162
 <211> 249
 <212> PRT
 <213> artificial

<220>
 <223> VH-VL of H2C

<400> 162

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
 100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 115 120 125

eo1f-seq1.txt

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 163
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL-P of H2C

<400> 163

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

eo1f-seq1.txt

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 164
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH-P of H2C

<400> 164

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 165
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL-P of H2C

<400> 165

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

eo1f-seq1.txt

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 166

eo1f-seq1.txt

<211> 109
<212> PRT
<213> artificial

<220>
<223> VL of H1E

<400> 166

Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 167
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH of H1E

<400> 167

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Glu Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

eo1f-seq1.txt

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Phe Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 168
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL of H1E

<400> 168

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Glu Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Phe Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
Page 143

eo1f-seq1.txt
155

145 150 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 169
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL-P of H1E

<400> 169

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

eo1f-seq1.txt

<210> 170
 <211> 125
 <212> PRT
 <213> artificial

<220>
 <223> VH-P of H1E

<400> 170

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Glu Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Phe Trp
 100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
 115 120 125

<210> 171
 <211> 249
 <212> PRT
 <213> artificial

<220>
 <223> VH-VL-P of H1E

<400> 171

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Glu Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val

eo1f-seq1.txt

35

40

45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Phe Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 172
<211> 125
<212> PRT
<213> artificial

<220>
<223> VL of G4H

eo1f-seq1.txt

<400> 172

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Arg Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Tyr Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 173

<211> 125

<212> PRT

<213> artificial

<220>

<223> VH of G4H

<400> 173

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Arg Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

eo1f-seq1.txt

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Tyr Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 174
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL of G4H

<400> 174

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Arg Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Tyr Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

eo1f-seq1.txt

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 175
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL-P of G4H

<400> 175

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 176

eof-seq1.txt

<211> 125
 <212> PRT
 <213> artificial

<220>
 <223> VH-P of G4H

<400> 176

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Arg Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Tyr Phe
 100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
 115 120 125

<210> 177
 <211> 249
 <212> PRT
 <213> artificial

<220>
 <223> VH-VL-P of G4H

<400> 177

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Arg Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

eo1f-seq1.txt

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Tyr Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 178
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL of A2J

<400> 178

Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly

eo1f-seq1.txt

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1              5              10              15
Thr Val Thr Leu Thr Cys Arg Ser Ser Thr Gly Ala Val Thr Ser Gly
      20      25      30
Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
      35      40      45
Leu Ile Gly Ala Thr Asp Met Arg Pro Ser Gly Thr Pro Ala Arg Phe
      50      55      60
Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
      65      70      75      80
Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
      85      90      95
Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
      100      105

<210> 179
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH of A2J

<400> 179
Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1      5      10      15
Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Val Tyr
      20      25      30
Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
      35      40      45
Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
      50      55      60
Ser Val Lys Lys Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
      65      70      75      80
Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
      85      90      95
Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Trp Trp
      100      105      110

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eo1f-seq1.txt

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 180
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL of A2J

<400> 180

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Val Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Lys Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Trp Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Arg Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Ala
180 185 190

eo1f-seq1.txt

Thr Asp Met Arg Pro Ser Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 181
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL-P of A2J

<400> 181

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Arg Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Ala Thr Asp Met Arg Pro Ser Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 182
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH-P of A2J

eo1f-seq1.txt

<400> 182

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Val Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Lys Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Trp Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 183

<211> 249

<212> PRT

<213> artificial

<220>

<223> VH-VL-P of A2J

<400> 183

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Val Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Lys Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

eo1f-seq1.txt

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Leu Ser Trp Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Arg Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Ala
180 185 190

Thr Asp Met Arg Pro Ser Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 184
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL of E1L

<400> 184

Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

eo1f-seq1.txt

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 185
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH of E1L

<400> 185

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Ser Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Thr Ser Tyr Tyr
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 186

eo1f-seq1.txt

<211> 249
 <212> PRT
 <213> artificial

<220>
 <223> VH-VL of E1L

<400> 186

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Ser Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Thr Ser Tyr Tyr
 100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
 130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
 165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
 180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
 195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
 210 215 220

eo1f-seq1.txt

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 187
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL-P of E1L

<400> 187

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 188
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH-P of E1L

<400> 188

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr

20

25

30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Ser Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Thr Ser Tyr Tyr
 100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
 115 120 125

<210> 189

<211> 249

<212> PRT

<213> artificial

<220>

<223> VH-VL-P of E1L

<400> 189

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Ser Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Thr Ser Tyr Tyr
 100 105 110

eo1f-seq1.txt

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 190
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL of E2M

<400> 190

Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Arg Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Ala Thr Asp Met Arg Pro Ser Gly Thr Pro Ala Arg Phe
50 55 60

eo1f-seq1.txt

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 191
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH of E2M

<400> 191

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Gly Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Glu Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Arg Asn Phe Gly Asn Ser Tyr Leu Ser Trp Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 192
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL of E2M

eo1f-seq1.txt

<400> 192

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15
Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Gly Tyr
20 25 30
Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45
Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60
Ser Val Lys Glu Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80
Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95
Tyr Cys Val Arg His Arg Asn Phe Gly Asn Ser Tyr Leu Ser Trp Phe
100 105 110
Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125
Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
130 135 140
Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160
Thr Cys Arg Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175
Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Ala
180 185 190
Thr Asp Met Arg Pro Ser Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205
Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220
Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240
Gly Gly Gly Thr Lys Leu Thr Val Leu

<210> 193
 <211> 109
 <212> PRT
 <213> artificial

<220>
 <223> VL-P of E2M
 <400> 193

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
 1 5 10 15

Thr Val Thr Leu Thr Cys Arg Ser Ser Thr Gly Ala Val Thr Ser Gly
 20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
 35 40 45

Leu Ile Gly Ala Thr Asp Met Arg Pro Ser Gly Thr Pro Ala Arg Phe
 50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
 65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
 85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
 100 105

<210> 194
 <211> 125
 <212> PRT
 <213> artificial

<220>
 <223> VH-P of E2M
 <400> 194

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Gly Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

eo1f-seq1.txt

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Glu Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Arg Asn Phe Gly Asn Ser Tyr Leu Ser Trp Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 195
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL-P of E2M
<400> 195

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Gly Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Glu Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Arg Asn Phe Gly Asn Ser Tyr Leu Ser Trp Phe
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
Page 165

eo1f-seq1.txt
140

130

135

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Arg Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Ala
180 185 190

Thr Asp Met Arg Pro Ser Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 196
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL of F70

<400> 196

Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

eo1f-seq1.txt

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 197
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH of F70

<400> 197

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Val Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Lys Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Trp Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 198
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL of F70

<400> 198

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Val Tyr

20

25

30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Lys Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Trp Trp
 100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
 130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
 165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
 180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
 195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
 210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
 225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
 245

<210> 199

<211> 109

<212> PRT

<213> artificial

eo1f-seq1.txt

<220>

<223> VL-P of F70

<400> 199

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Tyr Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 200

<211> 125

<212> PRT

<213> artificial

<220>

<223> VH-P of F70

<400> 200

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Val Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Lys Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

eo1f-seq1.txt

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Trp Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 201
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL-P of F70

<400> 201

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Val Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Lys Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Trp Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

eo1f-seq1.txt

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Tyr Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Ala Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 202
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL of F12Q

<400> 202

Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Asn Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 203

eo1f-seq1.txt

<211> 125
<212> PRT
<213> artificial

<220>
<223> VH of F12Q

<400> 203

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 204
<211> 249
<212> PRT
<213> artificial

<220>
<223> VH-VL of F12Q

<400> 204

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

eof-seq1.txt

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 205

<211> 109

<212> PRT

<213> artificial

<220>

<223> VL-P of F12Q

<400> 205

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly

eo1f-seq1.txt

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1              5              10              15
Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
      20      25      30
Asn Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
      35      40      45
Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
      50      55      60
Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
      65      70      75      80
Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn
      85      90      95
Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
      100      105

<210> 206
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH-P of F12Q

<400> 206
Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1      5      10
Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
      20      25      30
Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
      35      40      45
Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
      50      55      60
Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
      65      70      75      80
Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
      85      90      95
Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
      100      105      110

```

eo1f-seq1.txt

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 207

<211> 249

<212> PRT

<213> artificial

<220>

<223> VH-VL-P of F12Q

<400> 207

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Ser Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Val Ser Trp Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

eo1f-seq1.txt

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 208
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL of I2C

<400> 208

Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

Asn Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 209
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH of I2C

eo1f-seq1.txt

<400> 209

```

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1      5      10      15
Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
20      25      30
Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35      40      45
Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50      55      60
Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65      70      75      80
Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85      90      95
Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
100     105     110
Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115     120     125

```

<210> 210

<211> 249

<212> PRT

<213> artificial

<220>

<223> VH-VL of I2C

<400> 210

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Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1      5      10      15
Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
20      25      30
Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35      40      45
Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50      55      60
Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65      70      75      80

```

eo1f-seq1.txt

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
210 215 220

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 211
<211> 109
<212> PRT
<213> artificial

<220>
<223> VL-P of I2C

<400> 211

Glu Leu Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly
1 5 10 15

Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly
20 25 30

eo1f-seq1.txt

Asn Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly
35 40 45

Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe
50 55 60

Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val
65 70 75 80

Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn
85 90 95

Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
100 105

<210> 212
<211> 125
<212> PRT
<213> artificial

<220>
<223> VH-P of I2C

<400> 212

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
50 55 60

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser
115 120 125

<210> 213

eo1f-seq1.txt

<211> 249
 <212> PRT
 <213> artificial

<220>
 <223> VH-VL-P of I2C

<400> 213

Glu Val Gln Leu Leu Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
 20 25 30

Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 50 55 60

Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
 100 105 110

Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 115 120 125

Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Glu Leu Val Val
 130 135 140

Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 145 150 155 160

Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
 165 170 175

Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
 180 185 190

Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
 195 200 205

Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
 210 215 220

eo1f-seq1.txt

Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
225 230 235 240

Gly Gly Gly Thr Lys Leu Thr Val Leu
245

<210> 214
<211> 1587
<212> DNA
<213> artificial

<220>
<223> SA21-CD33 E11 HL x I2C HL (His Tag)

<400> 214
cgctgattg aagatatttg cctgccgcgc tggggctgcc tgtgggaaga tgatcagggtg 60
cagctggtgc agtctggagc tgaggatgaag aagcctggag agtcagtcaa ggtctcctgc 120
aaggctagcg ggtatacctt cacaaactat ggaatgaact gggatgaagca ggctccagga 180
cagggtttag agtggatggg ctggataaac acctacactg gagagccaac ctatgctgat 240
aagttccagg gacgcgttac catgactacg gatacctcta ccagcactgc ctatatggaa 300
atccgcaacc tcggagggtga tgacacggct gtatattact gtgcgcgctg gagttggagt 360
gatggttact acgtttactt tgactactgg ggccaaggca cttcggtcac cgtctcctca 420
gggtggtggtg gttctggcgg cggcggctcc ggtggtggtg gttctgacat cgtgatgaca 480
cagtctccag actccctgac tgtgtctctg ggcgagagga ccaccatcaa ctgcaagtcc 540
agccagagtg ttttagacag ctccacgaat aagaactcct tagcttggtg ccagcagaaa 600
ccaggacagc ctctctaaatt actcctttcc tgggcattcta cgcggaatc cgggatccct 660
gaccgattca gtggcagcgg gtctgggaca gatttcactc tcactattga cagcccgag 720
cctgaagatt ctgcaactta ctattgtcaa cagtctgccc acttcccgat cacctttggc 780
caagggacac gactggagat taaatccgga ggtggtggct ccgaggtgca gctggtcgag 840
tctggaggag gattggtgca gcctggaggg tcattgaaac tctcatgtgc agcctctgga 900
ttcaccttca ataagtacgc catgaactgg gtccgccagg ctccaggaaa gggtttgaa 960
tgggttgctc gcataagaag taaatataat aattatgcaa catattatgc cgattcagtg 1020
aaagacaggt tcaccatctc cagagatgat tcaaaaaaca ctgcctatct acaaatgaac 1080
aacttgaaaa ctgaggacac tgccgtgtac tactgtgtga gacatgggaa cttcggtaat 1140
agctacatat cctactgggc ttactggggc caagggactc tggtcaccgt ctctcagggt 1200
gggtggtggtt ctggcggcgg cggctccggt ggtggtggtt ctcagactgt tgtgactcag 1260
gaaccttcac tcaccgtatc acctgggtgga acagtcacac tcacttgtgg ctctcgact 1320

eof-seq1.txt

ggggctgtta catctggcaa ctacccaaac tgggtccaac aaaaaccagg tcaggcaccc 1380
 cgtgggtctaa taggtgggac taagttcctc gcccccggtta ctctgccag attctcaggc 1440
 tccctgcttg gaggaaggc tgccctcacc ctctcagggg tacagccaga ggatgaggca 1500
 gaatattact gtgttctatg gtacagcaac cgctgggtgt tcggtggagg aaccaaactg 1560
 actgtcctac atcaccatca ccatcac 1587

<210> 215
 <211> 529
 <212> PRT
 <213> artificial

<220>
 <223> SA21-CD33 E11 HL x I2C HL H6
 <400> 215

Arg Leu Ile Glu Asp Ile Cys Leu Pro Arg Trp Gly Cys Leu Trp Glu
 1 5 10 15

Asp Asp Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro
 20 25 30

Gly Glu Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr
 35 40 45

Asn Tyr Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu
 50 55 60

Trp Met Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp
 65 70 75 80

Lys Phe Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr
 85 90 95

Ala Tyr Met Glu Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr
 100 105 110

Tyr Cys Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp
 115 120 125

Tyr Trp Gly Gln Gly Thr Ser 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 Ser Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile
 165 170 175

eo1f-seq1.txt

Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn
 180 185 190
 Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu
 195 200 205
 Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser
 210 215 220
 Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Pro Gln
 225 230 235 240
 Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro
 245 250 255
 Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly
 260 265 270
 Gly Ser Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro
 275 280 285
 Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn
 290 295 300
 Lys Tyr Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu
 305 310 315 320
 Trp Val Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr
 325 330 335
 Ala Asp Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys
 340 345 350
 Asn Thr Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala
 355 360 365
 Val Tyr Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser
 370 375 380
 Tyr Trp Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly
 385 390 395 400
 Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr
 405 410 415
 Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val

420

425

430

Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr
 435 440 445

Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile
 450 455 460

Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly
 465 470 475 480

Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro
 485 490 495

Glu Asp Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp
 500 505 510

Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu His His His His His
 515 520 525

His

<210> 216
 <211> 1575
 <212> DNA
 <213> artificial

<220>
 <223> SA25-CD33 E11 HL x I2C HL with His tag

<400> 216
 gaggacatct gcctgcccag atggggctgc ctgtgggagg accaggtgca gctggtgcag 60
 tctggagctg aggtgaagaa gcctggagag tcagtcaagg tctcctgcaa ggctagcggg 120
 tataccttca caaactatgg aatgaactgg gtgaagcagg ctccaggaca gggtttagag 180
 tggatgggct ggataaacac ctacactgga gagccaacct atgctgataa gttccaggga 240
 cgcgttacca tgactacgga tacctctacc agcactgcct atatggaaat ccgcaacctc 300
 ggaggtgatg acacggctgt atattactgt gcgcgctgga gttggagtga tggttactac 360
 gtttactttg actactgggg ccaaggcact tcggtcaccg tctcctcagg tgggtggtgt 420
 tctggcggcg gcggctccgg tgggtggtgt tctgacatcg tgatgacaca gtctccagac 480
 tccctgactg tgtctctggg cgagaggacc accatcaact gcaagtccag ccagagtgtt 540
 ttagacagct ccacgaataa gaactcctta gcttgggtacc agcagaaacc aggacagcct 600
 cctaaattac tcctttcctg ggcattctac cggaatccg ggatccctga ccgattcagt 660
 ggcagcgggt ctgggacaga ttctactctc actattgaca gcccgcagcc tgaagattct 720

eof-seq1.txt

```

gcaacttact attgtcaaca gtctgcccac ttcccgatca cctttggcca agggacacga      780
ctggagatta aatccggagg tgggtggctcc gaggtgcagc tggtcgagtc tggaggagga      840
ttggtgcagc ctggaggggtc attgaaactc tcatgtgcag cctctggatt caccttcaat      900
aagtacgcca tgaactgggt ccgccaggct ccaggaaagg gtttggaatg ggttgctcgc      960
ataagaagta aatataataa ttatgcaaca tattatgccg attcagtgaag agacagggttc    1020
accatctcca gagatgattc aaaaaaact gcctatctac aaatgaacaa cttgaaaact      1080
gaggacactg ccgtgtacta ctgtgtgaga catgggaact tcggtaatag ctacatatcc      1140
tactgggctt actgggggcca agggactctg gtcaccgtct cctcagggtg tgggtggttct      1200
ggcggcgggcg gctccgggtg tgggtggttct cagactgttg tgactcagga accttcactc      1260
accgtatcac ctggtggaac agtcacactc acttgtggct cctcgactgg ggctgttaca      1320
tctggcaact acccaaactg ggtccaacaa aaaccaggtc aggcaccccg tgggtctaata      1380
ggtgggacta agttcctcgc ccccggtact cctgccagat tctcaggctc cctgcttggg      1440
ggcaaggctg ccctcacct ctcaggggta cagccagagg atgaggcaga atattactgt      1500
gttctatggg acagcaaccg ctgggtgttc ggtggaggaa ccaaactgac tgtcctacat      1560
caccatcacc atcac                                          1575

```

<210> 217
 <211> 525
 <212> PRT
 <213> artificial

<220>
 <223> SA25-CD33 E11 HL x I2C HL H6

<400> 217

Glu Asp Ile Cys Leu Pro Arg Trp Gly Cys Leu Trp Glu Asp Gln Val
 1 5 10 15

Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu Ser Val
 20 25 30

Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr Gly Met
 35 40 45

Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met Gly Trp
 50 55 60

Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe Gln Gly
 65 70 75 80

Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr Met Glu

Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr Tyr Cys Ala Arg
100 105 110

Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp Gly Gln
115 120 125

Gly Thr Ser Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly
130 135 140

Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser Pro Asp
145 150 155 160

Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys Lys Ser
165 170 175

Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu Ala Trp
180 185 190

Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser Trp Ala
195 200 205

Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser Gly Ser
210 215 220

Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Pro Gln Pro Glu Asp Ser
225 230 235 240

Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr Phe Gly
245 250 255

Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser Glu Val
260 265 270

Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly Ser Leu
275 280 285

Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr Ala Met
290 295 300

Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val Ala Arg
305 310 315 320

Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp Ser Val
325 330 335

eolf-seq1.txt

Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr Ala Tyr
340 345 350

Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr Tyr Cys
355 360 365

Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp Ala Tyr
370 375 380

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly Ser
385 390 395 400

Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val Thr Gln
405 410 415

Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu Thr Cys
420 425 430

Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn Trp Val
435 440 445

Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly Thr Lys
450 455 460

Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu Leu Gly
465 470 475 480

Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp Glu Ala
485 490 495

Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe Gly Gly
500 505 510

Gly Thr Lys Leu Thr Val Leu His His His His His His
515 520 525

<210> 218
<211> 1596
<212> DNA
<213> artificial

<220>
<223> SA08-CD33 E11 HL x I2C HL with His tag

<400> 218
cagggcctga tcggcgacat ctgcctgccc agatggggct gcctgtgggg cgactccgtg 60
aaacaggtgc agctggtgca gtctggagct gaggtgaaga agcctggaga gtcagtcaag 120
gtctcctgca aggctagcgg gtataccttc acaaactatg gaatgaactg ggtgaagcag 180

eolf-seq1.txt

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gctccaggac aggggtttaga gtggatgggc tggataaaca cctacactgg agagccaacc 240
tatgctgata agttccaggg acgcgttacc atgactacgg atacctctac cagcactgcc 300
tatatggaaa tccgcaacct cggaggtgat gacacggctg tatattactg tgcgcgctgg 360
agttggagtg atggttacta cgtttacttt gactactggg gccaaggcac ttcggtcacc 420
gtctcctcag gtggtggtgg ttctggcggc ggcggctccg gtggtggtgg ttctgacatc 480
gtgatgacac agtctccaga ctccctgact gtgtctctgg gcgagaggac caccatcaac 540
tgcaagtcca gccagagtgt tttagacagc tccacgaata agaactcctt agcttggtag 600
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gggatccctg accgattcag tggcagcggg tctgggacag atttcactct cactattgac 720
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acctttggcc aagggacacg actggagatt aaatccggag gtggtggctc cgaggtgcag 840
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tcctcgactg gggctgttac atctggcaac tacccaaaact gggccaaca aaaaccagg 1380
caggcacccc gtggtctaata aggtgggact aagttcctcg cccccgtac tcctgccaga 1440
ttctcaggct ccctgcttgg aggcaaggct gccctcacc tctcaggggt acagccagag 1500
gatgaggcag aatattactg tgttctatgg tacagcaacc gctgggtggt cggtggagga 1560
accaaactga ctgtcctaca tcaccatcac catcac 1596

```

<210> 219
 <211> 532
 <212> PRT
 <213> artificial

<220>
 <223> SA08-CD33 E11 HL x I2C HL H6

<400> 219

Gln Gly Leu Ile Gly Asp Ile Cys Leu Pro Arg Trp Gly Cys Leu Trp
 1 5 10 15

eo1f-seq1.txt

Gly Asp Ser Val₂₀ Lys Gln Val Gln₂₅ Leu Val Gln Ser Gly Ala₃₀ Glu Val

Lys Lys Pro₃₅ Gly Glu Ser Val₄₀ Lys Val Ser Cys Lys Ala₄₅ Ser Gly Tyr

Thr Phe₅₀ Thr Asn Tyr Gly Met₅₅ Asn Trp Val Lys Gln₆₀ Ala Pro Gly Gln

Gly₆₅ Leu Glu Trp Met Gly₇₀ Trp Ile Asn Thr Tyr₇₅ Thr Gly Glu Pro Thr₈₀

Tyr Ala Asp Lys Phe₈₅ Gln Gly Arg Val Thr₉₀ Met Thr Thr Asp Thr₉₅ Ser

Thr Ser Thr Ala₁₀₀ Tyr Met Glu Ile Arg₁₀₅ Asn Leu Gly Gly Asp₁₁₀ Asp Thr

Ala Val Tyr₁₁₅ Tyr Cys Ala Arg Trp₁₂₀ Ser Trp Ser Asp Gly₁₂₅ Tyr Tyr Val

Tyr Phe₁₃₀ Asp Tyr Trp Gly Gln₁₃₅ Gly Thr Ser Val Thr₁₄₀ Val Ser Ser Gly

Gly₁₄₅ Gly Gly Ser Gly Gly₁₅₀ Gly Gly Ser Gly Gly₁₅₅ Gly Gly Ser Asp Ile₁₆₀

Val Met Thr Gln₁₆₅ Ser Pro Asp Ser Leu Thr₁₇₀ Val Ser Leu Gly Glu₁₇₅ Arg

Thr Thr Ile Asn₁₈₀ Cys Lys Ser Ser Gln₁₈₅ Ser Val Leu Asp Ser₁₉₀ Ser Thr

Asn Lys Asn₁₉₅ Ser Leu Ala Trp Tyr₂₀₀ Gln Gln Lys Pro Gly₂₀₅ Gln Pro Pro

Lys Leu₂₁₀ Leu Leu Ser Trp Ala₂₁₅ Ser Thr Arg Glu Ser₂₂₀ Gly Ile Pro Asp

Arg₂₂₅ Phe Ser Gly Ser Gly₂₃₀ Ser Gly Thr Asp Phe₂₃₅ Thr Leu Thr Ile Asp₂₄₀

Ser Pro Gln Pro Glu₂₄₅ Asp Ser Ala Thr Tyr₂₅₀ Tyr Cys Gln Gln Ser₂₅₅ Ala

His Phe Pro Ile₂₆₀ Thr Phe Gly Gln Gly₂₆₅ Thr Arg Leu Glu Ile₂₇₀ Lys Ser

eo1f-seq1.txt

Gly Gly Gly Gly Ser Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu
275 280 285

Val Gln Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe
290 295 300

Thr Phe Asn Lys Tyr Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys
305 310 315 320

Gly Leu Glu Trp Val Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala
325 330 335

Thr Tyr Tyr Ala Asp Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp
340 345 350

Asp Ser Lys Asn Thr Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu
355 360 365

Asp Thr Ala Val Tyr Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser
370 375 380

Tyr Ile Ser Tyr Trp Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val
385 390 395 400

ser ser Gly Gly Gly Gly ser Gly Gly Gly Gly ser Gly Gly Gly Gly
405 410 415

ser Gln Thr Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly
420 425 430

Gly Thr Val Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser
435 440 445

Gly Asn Tyr Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg
450 455 460

Gly Leu Ile Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg
465 470 475 480

Phe Ser Gly Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly
485 490 495

Val Gln Pro Glu Asp Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser
500 505 510

Asn Arg Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu His His
515 520 525

eof-seq1.txt

His His His His
530

<210> 220
<211> 1569
<212> DNA
<213> artificial

<220>
<223> SA21-CD33 E11 HL x I2C HL

<400> 220
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cagctgggtgc agtctggagc tgagggtgaag aagcctggag agtcagtcaa ggtctcctgc 120
aaggctagcg ggtatacctt cacaactat ggaatgaact gggtagaagca ggctccagga 180
cagggtttag agtggatggg ctggataaac acctacactg gagagccaac ctatgctgat 240
aagttccagg gacgcgttac catgactacg gatacctcta ccagcactgc ctatatggaa 300
atccgcaacc tcggagggtga tgacacggct gtatattact gtgcgcgctg gaggttggagt 360
gatggttact acgtttactt tgactactgg ggccaaggca cttcggtcac cgtctcctca 420
ggtaggtggg gttctggcgg cggcggctcc ggtggtaggtg gttctgacat cgtgatgaca 480
cagctccag actccctgac tgtgtctctg ggcgagagga ccaccatcaa ctgcaagtcc 540
agccagagtg ttttagacag ctccacgaat aagaactcct tagcttggtg ccagcagaaa 600
ccaggacagc ctctaaatt actcctttcc tgggcatcta cgcgggaatc cgggatccct 660
gaccgattca gtggcagcgg gtctgggaca gatttcactc tactattga cagcccgcag 720
cctgaagatt ctgcaactta ctattgtcaa cagtctgccc acttcccgat cacctttggc 780
caagggacac gactggagat taaatccgga ggtggtaggtc ccgaggtgca gctggtcgag 840
tctggaggag gattgggtga gcctggaggg tcattgaaac tctcatgtgc agcctctgga 900
ttcaccttca ataagtacgc catgaactgg gtccgccagg ctccaggaaa gggtttgaa 960
tgggttgctc gcataagaag taaatataat aattatgcaa catattatgc cgattcagtg 1020
aaagacaggt tcaccatctc cagagatgat tcaaaaaaca ctgcctatct acaaatgaac 1080
aacttgaaaa ctgaggacac tgccgtgtac tactgtgtga gacatgggaa cttcggtaat 1140
agctacatat cctactgggc ttactggggc caagggactc tggtcaccgt ctctcaggt 1200
ggtaggtggt ctggcggcgg cggctccggt ggtggtaggt ctcagactgt tgtgactcag 1260
gaaccttcac tcaccgtatc acctgggtga acagtcacac tcaattgtgg ctctcgact 1320
ggggctgtta catctggcaa ctacccaaac tgggtccaac aaaaaccagg tcaggcaccc 1380
cgtggtctaa taggtgggac taagttcctc gccccggta ctctgccag attctcaggc 1440

eolf-seq1.txt

tccctgcttg gaggaaggc tgccctcacc ctctcagggg tacagccaga ggatgaggca 1500
 gaatattact gtgttctatg gtacagcaac cgctgggtgt tcggtggagg aaccaaactg 1560
 actgtccta 1569

<210> 221
 <211> 523
 <212> PRT
 <213> artificial

<220>
 <223> SA21-CD33 E11 HL x I2C HL

<400> 221

Arg Leu Ile Glu Asp Ile Cys Leu Pro Arg Trp Gly Cys Leu Trp Glu
 1 5 10 15

Asp Asp Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro
 20 25 30

Gly Glu Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr
 35 40 45

Asn Tyr Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu
 50 55 60

Trp Met Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp
 65 70 75 80

Lys Phe Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr
 85 90 95

Ala Tyr Met Glu Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr
 100 105 110

Tyr Cys Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp
 115 120 125

Tyr Trp Gly Gln Gly Thr Ser 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 Ser Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile
 165 170 175

Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn
 180 185 190

eo1f-seq1.txt

Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu
195 200 205

Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser
210 215 220

Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Pro Gln
225 230 235 240

Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro
245 250 255

Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly
260 265 270

Gly Ser Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro
275 280 285

Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn
290 295 300

Lys Tyr Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu
305 310 315 320

Trp Val Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr
325 330 335

Ala Asp Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys
340 345 350

Asn Thr Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala
355 360 365

Val Tyr Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser
370 375 380

Tyr Trp Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly
385 390 395 400

Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr
405 410 415

Val Val Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val
420 425 430

Thr Leu Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr

435

440

445

Pro Asn Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile
 450 455 460

Gly Gly Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly
 465 470 475 480

Ser Leu Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro
 485 490 495

Glu Asp Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp
 500 505 510

Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu
 515 520

<210> 222
 <211> 1557
 <212> DNA
 <213> artificial

<220>
 <223> SA25-CD33 E11 HL x I2C HL

<400> 222
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 tctggagctg aggtgaagaa gcctggagag tcagtcaagg tctcctgcaa ggctagcggg 120
 tataccttca caaactatgg aatgaactgg gtgaagcagg ctccaggaca gggtttagag 180
 tggatgggct ggataaacac ctacactgga gagccaacct atgctgataa gttccaggga 240
 cgcgttacca tgactacgga tacctctacc agcactgcct atatggaaat ccgcaacctc 300
 ggagggtgatg acacggctgt atattactgt gcgcgctgga gttggagtga tggttactac 360
 gtttactttg actactgggg ccaaggcact tcggtcaccg tctcctcagg tgggtggtggt 420
 tctggcggcg gcggctccgg tggtggtggt tctgacatcg tgatgacaca gtctccagac 480
 tccctgactg tgtctctggg cgagaggacc accatcaact gcaagtccag ccagagtgtt 540
 ttagacagct ccacgaataa gaactcctta gcttgggtacc agcagaaaacc aggacagcct 600
 cctaaattac tcctttcctg ggcatctacg cggaatccg ggatccctga ccgattcagt 660
 ggcagcgggt ctgggacaga ttctactctc actattgaca gccgcagcc tgaagattct 720
 gcaacttact attgtcaaca gtctgcccac ttcccgatca cctttggcca agggacacga 780
 ctggagatta aatccggagg tggtggtctc gaggtgcagc tggtcgagtc tggaggagga 840
 ttggtgcagc ctggaggggtc attgaaactc tcatgtgcag cctctggatt caccttcaat 900
 aagtacgcca tgaactgggt ccgccagggt ccaggaaaagg gtttggaatg ggttgctcgc 960

eof-seq1.txt

ataagaagta aatataataa ttatgcaaca tattatgccg attcagtgaa agacagggttc 1020
 accatctcca gagatgattc aaaaaaact gcctatctac aaatgaacaa cttgaaaact 1080
 gaggacactg ccgtgtacta ctgtgtgaga catgggaact tcggtaatag ctacatatcc 1140
 tactgggctt actggggcca agggactctg gtcaccgtct cctcagggtg tgggtggttct 1200
 ggcggcgccg gctccggtgg tgggtggttct cagactgttg tgactcagga accttcactc 1260
 accgtatcac ctggtggaac agtcacactc acttgtggct cctcgactgg ggctgttaca 1320
 tctggcaact acccaaactg ggtccaacaa aaaccagggtc aggcaccccg tgggtctaata 1380
 ggtgggacta agttcctcgc ccccggtact cctgccagat tctcaggctc cctgcttgga 1440
 ggcaaggctg ccctcaccct ctcaggggta cagccagagg atgaggcaga atattactgt 1500
 gttctatggt acagcaaccg ctgggtgttc ggtggaggaa ccaaactgac tgtccta 1557

<210> 223
 <211> 519
 <212> PRT
 <213> artificial

<220>
 <223> SA25-CD33 E11 HL x I2C HL

<400> 223

Glu Asp Ile Cys Leu Pro Arg Trp Gly Cys Leu Trp Glu Asp Gln Val
 1 5 10 15

Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Glu Ser Val
 20 25 30

Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Tyr Gly Met
 35 40 45

Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met Gly Trp
 50 55 60

Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe Gln Gly
 65 70 75 80

Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr Met Glu
 85 90 95

Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr Tyr Cys Ala Arg
 100 105 110

Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp Gly Gln
 115 120 125

eo1f-seq1.txt

Gly Thr Ser Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly Gly Gly
130 135 140

Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser Pro Asp
145 150 155 160

Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys Lys Ser
165 170 175

Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu Ala Trp
180 185 190

Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser Trp Ala
195 200 205

Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser Gly Ser
210 215 220

Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Pro Gln Pro Glu Asp Ser
225 230 235 240

Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr Phe Gly
245 250 255

Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser Glu Val
260 265 270

Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly Ser Leu
275 280 285

Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr Ala Met
290 295 300

Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val Ala Arg
305 310 315 320

Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp Ser Val
325 330 335

Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr Ala Tyr
340 345 350

Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr Tyr Cys
355 360 365

Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp Ala Tyr
370 375 380

eo1f-seq1.txt

Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly Gly Ser
385 390 395 400

Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val Thr Gln
405 410 415

Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu Thr Cys
420 425 430

Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn Trp Val
435 440 445

Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly Thr Lys
450 455 460

Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu Leu Gly
465 470 475 480

Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp Glu Ala
485 490 495

Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe Gly Gly
500 505 510

Gly Thr Lys Leu Thr Val Leu
515

<210> 224
<211> 1578
<212> DNA
<213> artificial

<220>
<223> SA08-CD33 E11 HL x I2C HL

<400> 224
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aaacaggtgc agctggtgca gtctggagct gaggtgaaga agcctggaga gtcagtcaag 120
gtctcctgca aggctagcgg gtataccttc acaaactatg gaatgaactg ggtgaagcag 180
gctccaggac aggggtttaga gtggatgggc tggataaaca cctacactgg agagccaacc 240
tatgctgata agttccaggg acgcgttacc atgactacgg atacctctac cagcactgcc 300
tatatggaaa tccgcaacct cggaggtgat gacacggctg tatattactg tgcgcgctgg 360
agttggagtg atggttacta cgtttacttt gactactggg gccaaggcac ttcggtcacc 420
gtctcctcag gtggtggtgg ttctggcggc ggcggctccg gtggtggtgg ttctgacatc 480

eof-seq1.txt

gtgatgacac agtctccaga ctccctgact gtgtctctgg gcgagaggac caccatcaac 540
 tgcaagtcca gccagagtgt tttagacagc tccacgaata agaactcctt agcttggtac 600
 cagcagaaac caggacagcc tcctaaatta ctcttttctt gggcatctac gcgggaatcc 660
 gggatccctg accgattcag tggcagcggg tctgggacag atttcaactct cactattgac 720
 agcccgagc ctgaagattc tgcaacttac tattgtcaac agtctgcca cttcccgatc 780
 acctttggcc aagggacacg actggagatt aaatccggag gtgggtggctc cgaggtgcag 840
 ctggtcgagt ctggaggagg attggtgcag cctggagggt cattgaaact ctcatgtgca 900
 gcctctggat tcaccttcaa taagtacgcc atgaactggg tccgccaggc tccaggaaag 960
 ggtttggaat ggggtgctcg cataagaagt aaatataata attatgcaac atattatgcc 1020
 gattcagtga aagacagggt caccatctcc agagatgatt caaaaaacac tgcctatcta 1080
 caaatgaaca acttgaaaac tgaggacact gccgtgtact actgtgtgag acatgggaac 1140
 ttcggtaata gctacatata ctactgggct tactggggcc aagggactct ggtcaccgtc 1200
 tcctcaggtg gtgggtggttc tggcggcggc ggctccggtg gtgggtggttc tcagactggt 1260
 gtgactcagg aaccttcaact caccgtatca cctgggtggaa cagtcacact cacttgtggc 1320
 tcctcgactg gggctgttac atctggcaac taccctaaact gggccaaca aaaaccagg 1380
 caggcacccc gtgggtctaata aggtgggact aagttcctcg ccccggtac tcctgccaga 1440
 ttctcaggct ccctgcttgg aggcaaggct gccctcacc tctcaggggt acagccagag 1500
 gatgaggcag aatattactg tgttctatgg tacagcaacc gctgggtggt cggtggagga 1560
 accaaactga ctgtccta 1578

<210> 225
 <211> 526
 <212> PRT
 <213> artificial

<220>
 <223> SA08-CD33 E11 HL x I2C HL
 <400> 225

Gln Gly Leu Ile Gly Asp Ile Cys Leu Pro Arg Trp Gly Cys Leu Trp
 1 5 10 15

Gly Asp Ser Val Lys Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val
 20 25 30

Lys Lys Pro Gly Glu Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr
 35 40 45

Thr Phe Thr Asn Tyr Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln
 50 55 60

eo1f-seq1.txt

Gly Leu Glu Trp Met Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr
 65 70 75 80
 Tyr Ala Asp Lys Phe Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser
 85 90 95
 Thr Ser Thr Ala Tyr Met Glu Ile Arg Asn Leu Gly Gly Asp Asp Thr
 100 105 110
 Ala Val Tyr Tyr Cys Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val
 115 120 125
 Tyr Phe 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 Asp Ile
 145 150 155 160
 Val Met Thr Gln Ser Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg
 165 170 175
 Thr Thr Ile Asn Cys Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr
 180 185 190
 Asn Lys Asn Ser Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro
 195 200 205
 Lys Leu Leu Leu Ser Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp
 210 215 220
 Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp
 225 230 235 240
 Ser Pro Gln Pro Glu Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala
 245 250 255
 His Phe Pro Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser
 260 265 270
 Gly Gly Gly Gly Ser Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu
 275 280 285
 Val Gln Pro Gly Gly Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe
 290 295 300
 Thr Phe Asn Lys Tyr Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys

eof-seq1.txt

```

<400> 226
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ccaggacagg gtttagagtg gatgggctgg ataaacacct aactggaga gccaacctat      180
gctgataagt tccagggacg cgttaccatg actacggata cctctaccag cactgcctat      240
atggaaaatcc gcaacctcgg aggtgatgac acggctgtat attactgtgc gcgctggagt      300
tggagtgatg gttactacgt ttactttgac tactggggcc aaggcacttc ggtcaccgtc      360
tcctcagggtg gtggtggttc tggcggcggc ggctccgggtg gtggtggttc tgacatcgtg      420
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aagtccagcc agagtgtttt agacagctcc acgaataaga actccttagc ttggtaccag      540
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eo1f-seq1.txt

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 20 25 30
 Gly Met Asn Trp Val Lys Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
 35 40 45
 Gly Trp Ile Asn Thr Tyr Thr Gly Glu Pro Thr Tyr Ala Asp Lys Phe
 50 55 60
 Gln Gly Arg Val Thr Met Thr Thr Asp Thr Ser Thr Ser Thr Ala Tyr
 65 70 75 80
 Met Glu Ile Arg Asn Leu Gly Gly Asp Asp Thr Ala Val Tyr Tyr Cys
 85 90 95
 Ala Arg Trp Ser Trp Ser Asp Gly Tyr Tyr Val Tyr Phe Asp Tyr Trp
 100 105 110
 Gly Gln Gly Thr Ser Val Thr Val Ser Ser Gly Gly Gly Gly Ser Gly
 115 120 125
 Gly Gly Gly Ser Gly Gly Gly Gly Ser Asp Ile Val Met Thr Gln Ser
 130 135 140
 Pro Asp Ser Leu Thr Val Ser Leu Gly Glu Arg Thr Thr Ile Asn Cys
 145 150 155 160
 Lys Ser Ser Gln Ser Val Leu Asp Ser Ser Thr Asn Lys Asn Ser Leu
 165 170 175
 Ala Trp Tyr Gln Gln Lys Pro Gly Gln Pro Pro Lys Leu Leu Leu Ser
 180 185 190
 Trp Ala Ser Thr Arg Glu Ser Gly Ile Pro Asp Arg Phe Ser Gly Ser
 195 200 205
 Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asp Ser Pro Gln Pro Glu
 210 215 220
 Asp Ser Ala Thr Tyr Tyr Cys Gln Gln Ser Ala His Phe Pro Ile Thr
 225 230 235 240
 Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Ser Gly Gly Gly Gly Ser
 245 250 255

eo1f-seq1.txt

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
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 Ser Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asn Lys Tyr
 275 280 285
 Ala Met Asn Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 290 295 300
 Ala Arg Ile Arg Ser Lys Tyr Asn Asn Tyr Ala Thr Tyr Tyr Ala Asp
 305 310 315 320
 Ser Val Lys Asp Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 325 330 335
 Ala Tyr Leu Gln Met Asn Asn Leu Lys Thr Glu Asp Thr Ala Val Tyr
 340 345 350
 Tyr Cys Val Arg His Gly Asn Phe Gly Asn Ser Tyr Ile Ser Tyr Trp
 355 360 365
 Ala Tyr Trp Gly Gln Gly Thr Leu Val Thr Val Ser Ser Gly Gly Gly
 370 375 380
 Gly Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gln Thr Val Val
 385 390 395 400
 Thr Gln Glu Pro Ser Leu Thr Val Ser Pro Gly Gly Thr Val Thr Leu
 405 410 415
 Thr Cys Gly Ser Ser Thr Gly Ala Val Thr Ser Gly Asn Tyr Pro Asn
 420 425 430
 Trp Val Gln Gln Lys Pro Gly Gln Ala Pro Arg Gly Leu Ile Gly Gly
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 Thr Lys Phe Leu Ala Pro Gly Thr Pro Ala Arg Phe Ser Gly Ser Leu
 450 455 460
 Leu Gly Gly Lys Ala Ala Leu Thr Leu Ser Gly Val Gln Pro Glu Asp
 465 470 475 480
 Glu Ala Glu Tyr Tyr Cys Val Leu Trp Tyr Ser Asn Arg Trp Val Phe
 485 490 495
 Gly Gly Gly Thr Lys Leu Thr Val Leu His His His His His His

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