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(54) Title: PEPTIDE SCAFFOLD DESIGN

X^1 - X^2 - X^3 - X^4 - X^5 (I)

(57) Abstract: The present invention relates to novel peptides and methods for treatment, diagnosis and prognosis of virus infections including infections with HCV, HIV, CMV and Influenza. The invention further relates to methods for identifying and providing peptides useful for the treatment and diagnosis.

PEPTIDE SCAFFOLD DESIGN

FIELD OF THE INVENTION

The present invention relates to novel peptides and methods for treatment, diagnosis and prognosis of virus infections including infections with HCV, HIV, CMV and Influenza. The invention further relates to methods for identifying and providing peptides useful for the treatment and diagnosis.

BACKGROUND OF THE INVENTION

Conventional approaches to vaccine development have implemented either whole replication competent virus which has been attenuated (e.g. Sabin polio vaccine, measles, mumps, rubella (MMR)) or inactivated virions that are not replication competent. On occasions, the inactivated virus vaccines may include split vaccines where the virus particles have been disrupted. Molecular techniques have also been used to develop the subunit vaccine (e.g. hepatitis B vaccine) that consists only of the surface glycoproteins of hepatitis B virus. The inactivated virus vaccines tend to induce primarily antibody responses to the viruses in question, whereas the live attenuated vaccines induce both cell-mediated immunity as well as an antibody response since the vaccine induces a transient infection.

The only disease which has been eliminated by virtue of a successful vaccination campaign is smallpox. A campaign is currently in progress to eradicate polio. Features of virus infections that can be eliminated by vaccination are infections caused by viruses with stable virus antigens (i.e. very low mutation frequency, few subtypes), that lack a reservoir in other animal species, viruses that do not persist in the body once the infection is over and where vaccination leads to long lasting immunity. Viruses such as polio and measles fulfill these criteria whereas viruses such as influenza virus (Flu), HCV, and HIV that vary their protein sequences do not. It is for this reason that new and alternate approaches are required to develop vaccines for these diseases.

Vaccination aims to stimulate the immune response to a specific pathogen in advance of infection. When an individual is exposed to that pathogen, a memory response is triggered which prevents the establishment of infection. Vaccines therefore stimulate the adaptive immune response which unlike innate immunity, is long lived and has memory. There are two major arms to the adaptive immune system. Humoral immunity which involves the development of antibodies that can bind virus particles and certain antibodies that can neutralize infection. Cell mediated immunity that leads to the development of cytotoxic T-

cells that kill infected cells exposing viral epitopes in the context of human leukocyte antigen (HLA) class I, in this way eliminating infected cells.

The challenge of providing vaccines suitable for stimulation of the adaptive immune system is that peptide epitopes need to be taken up by the antigen presenting cells.

- 5 Several peptides have been demonstrated to translocate across the plasma membrane of eukaryotic cells by a seemingly energy- independent pathway. These peptides are defined as cell-penetrating peptides (CPPs). Cellular delivery using these cell-penetrating peptides offers several advantages over conventional techniques. It is non-invasive, energy-independent, is efficient for a broad range of cell types and can be applied to cells en masse.
- 10 Hepatitis means inflammation of the liver which can be caused by a variety of factors including toxins, certain drugs, some diseases, heavy alcohol use, and bacterial and viral infections. Hepatitis is also the name of a family of viral infections that affect the liver; the most common types in the developed world are hepatitis A, hepatitis B, and hepatitis C.

- Hepatitis C is a liver disease that results from infection with the hepatitis C virus (HCV). It can range in severity from a mild illness lasting a few weeks to a serious, lifelong illness. Hepatitis C is spread via blood; the most common form of transmission is through sharing needles or other equipment used to inject drugs. The infection can be either "acute" or "chronic". Acute HCV infection is an asymptomatic, short-term illness that occurs within the first 6 months after someone is exposed to the hepatitis C virus. For most people, acute infection leads to chronic infection, which can result in long-term complications and even death.
- 15
- 20

- HCV is an enveloped positive stranded ribonucleic acid (RNA) virus with a diameter of about 50nm, belonging to the genus Hepacivirus in the family Flaviviridae that replicate in the cytoplasm of infected cells. The only known reservoir for HCV is humans, although the virus has experimentally been transmitted to chimpanzees. The natural targets of HCV are hepatocytes and possibly B-lymphocytes. As of 2008, six different genotypes and more than 100 subtypes of the virus are known. Replication occurs through an RNA-dependent RNA polymerase that lacks a proofreading function, which results in a very high rate of mutations. Rapid mutations in a hypervariable region of the HCV genome coding for the envelope proteins enable the virus to escape immune surveillance by the host. As a consequence, most HCV-infected people proceed to chronic infection.
- 25
- 30

It is estimated that 170 million people are infected with HCV worldwide, equating to approximately 3% of the global population. There are also approximately 3-4 million people

who are infected every year; with an estimated 80% of these newly infected patients progressing to chronic infection.

The 6 genotypes of HCV have different geographical spread. The disease in the early stages is generally asymptomatic; the majority of patients with chronic infection eventually progress to complications such as liver fibrosis and cirrhosis, and, in 1-5% of cases, hepatocellular carcinoma.

HCV is the major cause of non-A, non-B hepatitis worldwide. Acute infection with HCV frequently leads to chronic hepatitis and end-stage cirrhosis. It is estimated that up to 20% of HCV chronic carriers may develop cirrhosis over a time period of about 20 years and that of those with cirrhosis between 1 to 4% is at risk to develop liver carcinoma.

The about 9.6 kb single-stranded RNA genome of the HCV virus comprises a 5'- and 3'-noncoding region (NCRs) and, in between these NCRs a single long open reading frame of about 9 kb encoding an HCV polypeptide of about 3000 amino acids.

HCV polypeptides are produced by translation from the open reading frame and cotranslational proteolytic processing. Structural proteins are derived from the amino-terminal one-fourth of the coding region and include the capsid or Core protein (about 21 kDa), the E1 envelope glycoprotein (about 35 kDa) and the E2 envelope glycoprotein (about 70 kDa, previously called NS1), and p7 (about 7kDa). The E2 protein can occur with or without a C-terminal fusion of the p7 protein (Shimotohno et al. 1995). An alternative open reading frame in the Core-region has been found which is encoding and expressing a protein of about 17 kDa called F (Frameshift) protein (Xu et al. 2001; Ou & Xu in US Patent Application Publication No. US2002/0076415). In the same region, ORFs for other 14-17 kDa ARFPs (Alternative Reading Frame Proteins), A1 to A4, were discovered and antibodies to at least A1, A2 and A3 were detected in sera of chronically infected patients (Walewski et al. 2001). From the remainder of the HCV coding region, the non-structural HCV proteins are derived which include NS2 (about 23 kDa), NS3 (about 70 kDa), NS4A (about 8 kDa), NS4B (about 27 kDa), NS5A (about 58 kDa) and NS5B (about 68 kDa) (Grakoui et al. 1993).

Influenza remains a significant cause of mortality and morbidity worldwide. The World Health Organisation (WHO) estimates that seasonal epidemics affect 3-5 million people with severe illness annually and result in 250,000 – 500,000 mortalities. Influenza is caused by viruses in the family Orthomyxoviridae which are negative stranded RNA viruses. The influenza virus exists as three types, A, B and C of which only A is associated with pandemics. Types A viruses are found in both humans and animals, particularly birds but also other mammals such as pigs. Type A viruses are further typed into subtypes according to different kinds and

combinations of virus surface proteins. Among many subtypes in 2009 influenza A (H1N1) and A (H3N2) subtypes were circulating among humans. Influenza A and B are included in the seasonal vaccine, whereas influenza C occurs only rarely, and so it is not included in the seasonal vaccine. Type B viruses are human specific and Type C viruses cause a very mild disease. The genomes of Orthomyxoviruses are segmented. Influenza viruses Types A and B have 8 segments whereas type C has seven. Pandemics may arise as a result of re-assortment of gene segments when two different type A viruses infect the same cell. There is no immunity in the population to this novel re-assorted virus. Three pandemics occurred in the twentieth century: "Spanish influenza" in 1918, "Asian influenza" in 1957, and "Hong Kong influenza" in 1968. The 1918 pandemic killed an estimated 40–50 million people worldwide. Subsequent pandemics were much milder, with an estimated 2 million deaths in 1957 and 1 million deaths in 1968. In June 2009 the WHO declared a pandemic from influenza virus H1N1 (swine Influenza) which was declared over in August 2010.

Human papillomaviruses are made up of a group of DNA viruses in the family Papillomaviridae which infect the skin and mucous membranes. Two groups which are derived from more than 100 different identified subtypes are the main cause for clinical concern: those causing warts (both benign and genital warts), and a group of 12 "high risk" subtypes that can result in cervical cancer. This latter group has been attributed as a contributory factor in the development of nearly all types of cervical cancer. Worldwide, cervical cancer remains the second most common malignancy in women, and is a leading cause of cancer-related death for females in developing countries. HPV 16 and 18 have been mainly associated with cervical cancer, however, the virus is also a cause of throat cancer in both men and women. HPV is transmitted through contact and enters the skin through abrasions. An abortive infection, where only the early proteins are expressed is associated with cancer development.

OVERVIEW OF THE INVENTION

The invention provides peptides that may be used as immunogens to stimulate an adaptive immune response in a subject.

The invention also provides peptides that may be taken up by antigen presenting cells (macrophages and dendritic cells) such that epitopes within the peptides are correctly processed and presented to T-lymphocytes in order to stimulate an effective immune response.

The invention further provides peptides that may be used as antigens, to provide immunogenic compositions and methods for inducing an immune response in a subject against an antigen.

SUMMARY OF THE INVENTION

5 The present invention pertains to a peptide design promoting uptake of peptide epitopes by antigen presenting cells (macrophages and dendritic cells) such that the epitopes can be correctly processed and presented in the context of HLA class I and II to stimulate both CD4+ and CD8+ T-lymphocytes. CD8+ T-lymphocytes with cytotoxic capacity will kill infected cells bearing the epitope of interest. CD4+ T-lymphocyte provide 'help' to sustain
10 effective CD8+ T-lymphocyte responses.

It has been found by the present inventor(s) that peptide constructs - amino acid sequences with a particular pattern or scaffold design - have the ability to effectively penetrate the cell membrane. Accordingly, the peptide constructs according to the present invention may be used to load cells with an immunogenically effective amount of a peptide
15 or fragments of this peptide that can be presented by macrophages and dendritic cells. Accordingly these peptide constructs may elicit a Cytotoxic T-lymphocyte immune (CTL) response and/or a Humoral Immune Response.

So, in a first aspect the present invention relates to an isolated cell-penetrating peptide of not more than 60 amino acids with the following structure

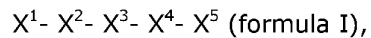
20 $X^1 - X^2 - X^3 - X^4 - X^5$ (formula I),

wherein X^1 and X^3 independently defines a linear sequence of any 1, 2, 3 or 4 amino acid(s) independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof; X^2 defines a linear sequence of 8-30 amino acids derived from an antigen; X^4 defines a linear sequence of 8-30 amino acids derived from said antigen, said
25 sequence X^4 being different from X^2 ; and wherein X^5 is any one optional amino acid selected from a basic amino acid, citrulline, tryptophan, or a derivative thereof.

It is to be understood that the amino acid sequence of formula I refers to a peptide sequence in a standard N- to C-terminal direction, wherein the first amino acid mentioned is the N-terminal amino acid that may have an amino ($-NH_2$) group or alternatively an -
30 NH_3^+ group. The last amino acid mentioned is the C-terminal that may have a free carboxyl group ($-COOH$) or a carboxylate group. In some embodiments the N- and/or C-terminal amino acid is

modified, such as by N-terminal acetylation or C-terminal amidation. The symbol "-" used in formula I refers to a standard peptide bond, such as a standard peptide bond between X^1 and X^2 in " X^1 - X^2 ".

It is further to be understood that the peptides according to the invention primarily are intended for synthetic peptide synthesis. However, the peptides may be longer than 60 amino acids, if the peptides are produced by recombinant means. Accordingly, an alternative first aspect of the present invention relates to an isolated cell-penetrating peptide comprising a peptide sequence with the following structure



wherein X^1 and X^3 independently defines a linear sequence of any 1, 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof; X^2 defines a linear sequence of 8-30 amino acids derived from an antigen; X^4 defines a linear sequence of 8-30 amino acids derived from said antigen, said sequence X^4 being different from X^2 ; and wherein X^5 is any one optional amino acid selected from a basic amino acid, citrulline, tryptophan, or a derivative thereof.

In a second aspect the present invention relates to an isolated peptide of not more than 60 amino acids comprising a sequence of X^2 and/or X^4 as independently defined in any one of table 1 or table 2.

In a third aspect the present invention relates to the use of a peptide comprising a sequence of X^2 or X^4 as independently defined in any one of table 1 or table 2 for inducing an immune response in a subject.

In a further aspect the present invention relates to an isolated peptide consisting of $X^1 - X^5$ as defined in any one of table 1 or table 2.

In a further aspect the present invention relates to a dimer peptide comprising two peptide monomers, wherein each peptide monomer is according to the invention.

In a further aspect the present invention relates to a peptide combination comprising two or more peptides according to the invention.

In a further aspect the present invention relates to an isolated nucleic acid or polynucleotide encoding a peptide according to the invention.

In a further aspect the present invention relates to a vector comprising the nucleic acid or polynucleotide encoding a peptide according to the invention.

In a further aspect the present invention relates to a host cell comprising the vector comprising the nucleic acid or polynucleotide encoding a peptide according to the invention.

- 5 In a further aspect the present invention relates to an immunogenic composition comprising at least one peptide according to the invention, a dimer peptide comprising two peptides monomers, wherein each peptide monomer is according to the invention, a peptide combination comprising two or more peptide according to the invention, the nucleic acid or polynucleotide encoding a peptide according to the invention, or the vector comprising the
10 nucleic acid or polynucleotide encoding a peptide according to the invention; in combination with a pharmaceutically acceptable diluent or vehicle and optionally an immunological adjuvant.

- In a further aspect the present invention relates to a method for inducing an immune response in a subject against an antigen which comprises administration of at least one
15 peptide according to the invention; a dimer peptide comprising two peptides monomers, wherein each peptide monomer is according to the invention; a peptide combination comprising two or more peptide according to the invention; the nucleic acid or polynucleotide encoding a peptide according to the invention; the vector comprising the nucleic acid or polynucleotide encoding a peptide according to the invention; or the composition according to
20 the invention.

- In a further aspect the present invention relates to a method for reducing and/or delaying the pathological effects of a virus in a subject infected with said virus, the method comprising administering an effective amount of at least one peptide according to the invention; a dimer peptide comprising two peptides monomers, wherein each peptide monomer is according to
25 the invention; a peptide combination comprising two or more peptide according to the invention; the nucleic acid or polynucleotide encoding a peptide according to the invention; the vector comprising the nucleic acid or polynucleotide encoding a peptide according to the invention; or the composition according to the invention.

- In a further aspect the present invention relates to a peptide according to the invention for
30 use as a medicament.

In a further aspect the present invention relates to a peptide according to the invention for treating the pathological effects of a virus in a subject infected with said virus.

In a further aspect the present invention relates to an isolated cell-penetrating peptide consisting of the following structure

X^1 - X^2 - X^3 - X^4 - X^5 (formula I),

wherein:

5 X^1 and X^3 independently define a linear sequence of any 2, or 3 arginines;

X^1 and/or X^3 consist of RR or RRR;

X^2 and X^4 independently define a linear sequence of 8-15 amino acids having at least 70% sequence identity to an antigen from HCV, CMV, HPV, or Influenza or a variant sequence thereof;

10 the sequence of X^4 is derived from the same antigen as X^2 but is different from the sequence of X^2 ;

X^5 is an optional arginine;

wherein the sequence of amino acids defined by X^1 - X^2 - X^3 - X^4 - X^5 of formula I is not found in the native sequence of said antigen; and

15 wherein the sequence of amino acids defined by X^1 - X^2 - X^3 - X^4 - X^5 of formula I does not contain a continuous sequence of four or more arginine residues.

Throughout this specification the word "comprise", or variations such as "comprises" or "comprising", will be understood to imply the inclusion of a stated element, integer or step,
20 or group of elements, integers or steps, but not the exclusion of any other element, integer or step, or group of elements, integers or steps.

Any discussion of documents, acts, materials, devices, articles or the like which has been included in the present specification is not to be taken as an admission that any or all of
25 these matters form part of the prior art base or were common general knowledge in the field relevant to the present disclosure as it existed before the priority date of each claim of this application.

DETAILED DISCLOSURE OF THE INVENTION

30 *Definitions*

When terms such as "one", "a" or "an" are used in this disclosure they mean "at least one", or "one or more" unless otherwise indicated. Further, the term "comprising" is intended to mean "including" and thus allows for the presence of other constituents, features, conditions, or steps than those explicitly recited.

"HIV" generally denotes human immunodeficiency virus I.

"HIV disease" is composed of several stages including the acute HIV infection which often manifests itself as an influenza-like infection and the early and medium stage symptomatic disease, which has several non-characteristic symptoms such as skin rashes, fatigue, night sweats, slight weight loss, mouth ulcers, and fungal skin and nail infections. Most HIV infected will experience mild symptoms such as these before developing more serious illnesses. It is generally believed that it takes five to seven years for the first mild symptoms to appear. As HIV disease progresses, some individuals may become quite ill even if they have not yet been diagnosed with AIDS (see below), the late stage of HIV disease. Typical problems include chronic oral or vaginal thrush (a fungal rash or spots), recurrent herpes blisters on the mouth (cold sores) or genitals, ongoing fevers, persistent diarrhea, and significant weight loss. "AIDS" is the late stage HIV disease and is a condition which progressively reduces the effectiveness of the immune system and leaves individuals susceptible to opportunistic infections and tumors.

The term "cell-penetrating peptide" as used herein refers to any peptide with the capability to translocate across the plasma membrane into either cytoplasmic and/or nuclear compartments of eukaryotic and/or prokaryotic cells, such as into cytoplasm, nucleus, lysosome, endoplasmatic reticulum, golgi apparatus, mitochondria and/or chloroplast, seemingly energy-independently. This capability to translocate across the plasma membrane of a "cell-penetrating peptide" according to the invention may be non-invasive, energy-independent, non-saturable, and/or receptor independent. In one embodiment the term "cell-penetrating peptide" refers to a peptide, which is demonstrated to translocate across a plasma membrane as determined by the assay in example 1. It is to be understood that a cell-penetrating peptide according to the present invention may be translocated across the membrane with the sequence complete and intact, or alternatively partly degraded, but in a form where the antigens contained within this peptide is able to be presented within the cell to stimulate an immune response. Accordingly, a cell-penetrating peptide according to the

present invention is a peptide that may be demonstrated to translocate across a plasma membrane as determined by the assay in example 1 and be demonstrated to stimulate an effective immune response.

The term "derived from an antigen" when in reference to a peptide derived from a source (such as a virus etc.) as used herein is intended to refer to a peptide which has been obtained (e.g., isolated, purified, etc.) from the source. Preferably, the peptide may be genetically engineered and/or chemically synthesized to be essentially identical to the native peptide of the source. The term includes the use of variants of known native peptide sequences, such as peptide sequences, where 1, 2, 3, 4, 5, 6, 7, 8, 9, or 10 amino acids of the native peptide sequence have been substituted with any other amino acid, such as conservative substitutions. Alternatively, 1, 2, 3, 4, 5, 6, 7, 8, 9, or 10 amino acids have been removed or added to the native peptide sequence. Accordingly, in some embodiments, the peptides according to the present invention comprises the sequences X^2 and/or X^4 , that is defined as a sequence of 8-30 amino acids, such as 8-20 amino acids derived from an antigen, wherein the peptide sequence of the antigen comprises 1, 2, 3, 4, 5, 6, 7, 8, 9, or 10 substitutions, additions or deletions relative to the antigen, such as the addition of an arginine in the N- or C-terminal of the amino acid sequence of X^2 and/or X^4 . The amino acids used in the amino acid sequences according to the invention may be in both L- and/or D-form. It is to be understood that both L- and D-forms may be used for different amino acids within the same peptide sequence. In some embodiments the amino acids within the peptide sequence are in L-form, such as natural amino acids. It is to be understood that any known antigen may be used in the constructs according to the present invention.

In some specific embodiments, the first 1, 2, or 3 amino acids in the N-terminal of the amino acid sequences according to the invention are in the D-form. It is assumed that the N-terminal trimming and thereby degradation of the peptides are somewhat delayed by having amino acids of the D-form in the N-terminal of these cell-penetrating peptides. Alternatively and in some embodiments, the first 1, 2, or 3 amino acids in the N-terminal of the amino acid sequences according to the invention are amino acids in beta or gamma forms. Beta amino acids have their amino group bonded to the beta carbon rather than the alpha carbon as in the 20 standard natural amino acids.

Alternatively the first 1, 2, or 3 amino acids in the N-terminal of the amino acid sequences according to the invention may be modified by incorporation of fluorine, or alternatively cyclic amino acids or other suitable non-natural amino acids are used.

A "variant" or "analogue" of a peptide refers to a peptide having an amino acid sequence that is substantially identical to a reference peptide, typically a native or "parent" polypeptide. The

peptide variant may possess one or more amino acid substitutions, deletions, and/or insertions at certain positions within the native amino acid sequence.

"Conservative" amino acid substitutions are those in which an amino acid residue is replaced with an amino acid residue having a side chain with similar physicochemical properties.

- 5 Families of amino acid residues having similar side chains are known in the art, and include amino acids with basic side chains (e.g., lysine, arginine, histidine), acidic side chains (e.g., aspartic acid, glutamic acid), uncharged polar side chains (e.g., glycine, asparagine, glutamine, serine, threonine, tyrosine, cysteine, tryptophan), nonpolar side chains (e.g., alanine, valine, leucine, isoleucine, proline, phenylalanine, methionine), beta-branched side
- 10 chains (e.g., threonine, valine, isoleucine) and aromatic side chains (e.g., tyrosine, phenylalanine, tryptophan, histidine). A particular form of conservative amino acid substitutions include those with amino acids, which are not among the normal 20 amino acids encoded by the genetic code. Since preferred embodiments of the present invention entail use of synthetic peptides, it is unproblematic to provide such "non-naturally occurring" amino
- 15 acid residues in the peptides disclosed herein, and thereby it is possible to exchange the natural saturated carbon chains in the side chains of amino acid residues with shorter or longer saturated carbon chains – for instance, lysine may be substituted with an amino acid having an the side chain $-(CH_2)_nNH_3$, where n is different from 4, and arginine may be substituted with an amino acid having the side chain $-(CH_2)_nNHC(=NH_2)NH_2$, where n is
- 20 different from 3, etc. Similarly, the acidic amino acids aspartic acid and glutamic acid may be substituted with amino acid residues having the side chains $-(CH_2)_nCOOH$, where $n > 2$.

- The term "substantially identical" in the context of two amino acid sequences means that the sequences, when optimally aligned, such as by the programs GAP or BESTFIT using default gap weights, share at least about 50, at least about 60, at least about 70, at least about 80,
- 25 at least about 90, at least about 95, at least about 98, or at least about 99 percent sequence identity. In one embodiment, residue positions that are not identical differ by conservative amino acid substitutions. Sequence identity is typically measured using sequence analysis software. Protein analysis software matches similar sequences using measures of similarity assigned to various substitutions, deletions and other modifications, including conservative
- 30 amino acid substitutions. For instance, the publicly available GCG software contains programs such as "Gap" and "BestFit" which can be used with default parameters to determine sequence homology or sequence identity between closely related polypeptides, such as homologous polypeptides from different species of organisms or between a wild-type protein and a mutein thereof. See, *e.g.*, GCG Version 6.1. Polypeptide sequences can also be
- 35 compared using FASTA or ClustalW, applying default or recommended parameters. A program in GCG Version 6.1., FASTA (*e.g.*, FASTA2 and FASTA3) provides alignments and percent sequence identity of the regions of the best overlap between the query and search

sequences (Pearson, Methods Enzymol. 1990;183:63-98; Pearson, Methods Mol. Biol. 2000;132:185-219). Another preferred algorithm when comparing a sequence to a database containing a large number of sequences from various organisms, or when deducing the is the computer program BLAST, especially blastp, using default parameters. See, *e.g.*, Altschul et al., J. Mol. Biol. 1990;215:403-410; Altschul et al., Nucleic Acids Res. 1997;25:3389-402 (1997); each herein incorporated by reference. "Corresponding" amino acid positions in two substantially identical amino acid sequences are those aligned by any of the protein analysis software mentioned herein, typically using default parameters.

An "isolated" molecule is a molecule that is the predominant species in the composition wherein it is found with respect to the class of molecules to which it belongs (*i.e.*, it makes up at least about 50% of the type of molecule in the composition and typically will make up at least about 70%, at least about 80%, at least about 85%, at least about 90%, at least about 95%, or more of the species of molecule, *e.g.*, peptide, in the composition). Commonly, a composition of a peptide molecule will exhibit 98% - 99% homogeneity for peptide molecules in the context of all present peptide species in the composition or at least with respect to substantially active peptide species in the context of proposed use.

The term "linear sequence" as used herein refers to the specific sequence of amino acids connected by standard peptide bonds in standard N- to C-terminal direction. The peptide may contain only peptide bonds. However the term does not exclude that an amino acid within a sequence, such as within X³, may be connected, such as through the side chains, with another amino acid at a distant location within the peptide sequence, such as a distant location within X³.

In the context of the present invention, "treatment" or "treating" refers to preventing, alleviating, managing, curing or reducing one or more symptoms or clinically relevant manifestations of a disease or disorder, unless contradicted by context. For example, "treatment" of a patient in whom no symptoms or clinically relevant manifestations of a disease or disorder have been identified is preventive or prophylactic therapy, whereas "treatment" of a patient in whom symptoms or clinically relevant manifestations of a disease or disorder have been identified generally does not constitute preventive or prophylactic therapy.

The term "antigen" denotes a substance of matter which is recognized by the immune system's specifically recognizing components (antibodies, T-cells).

The term "immunogen" is in the present context intended to denote a substance of matter, which is capable of inducing an adaptive immune response in an individual, where said

adaptive immune response targets the immunogen. In relation to the present invention, an immunogen will induce a humoral and/or cell-mediated immune response. In other words, an immunogen is an antigen, which is capable of inducing immunity.

5 The terms "epitope", "antigenic determinant" and "antigenic site" are used interchangeably herein and denotes the region in an antigen or immunogen which is recognized by antibodies (in the case of antibody binding epitopes, also known as "B-cell epitopes") or by T-cell receptors when the epitope is complexed to a Major histocompatibility complex (MHC) molecule (in the case of T-cell receptor binding epitopes, i.e. "T-cell epitopes").

10 The term "immunogenically effective amount" has its usual meaning in the art, *i.e.* an amount of an immunogen, which is capable of inducing an immune response, which significantly engages pathogenic agents, which share immunological features with the immunogen.

15 The term "vaccine" is used for a composition comprising an immunogen and which is capable of inducing an immune response which is either capable of reducing the risk of developing a pathological condition or capable of inducing a therapeutically effective immune response which may aid in the cure of (or at least alleviate the symptoms of) a pathological condition.

20 The term "pharmaceutically acceptable" has its usual meaning in the art, *i.e.* it is used for a substance that can be accepted as part of a medicament for human use when treating the disease in question and thus the term effectively excludes the use of highly toxic substances that would worsen rather than improve the treated subject's condition.

25 A "T helper lymphocyte epitope" (a T_H epitope) is peptide, which binds an MHC Class II molecule and can be presented on the surface of an antigen presenting cell (APC) bound to the MHC Class II molecule. An "immunological carrier" is generally a substance of matter which includes one or many T_H epitopes, and which increase the immune response against an antigen to which it is coupled by ensuring that T-helper lymphocytes are activated and proliferate. Examples of known immunological carriers are the tetanus and diphtheria toxoids and keyhole limpet hemocyanin (KLH).

30 In the scaffold design according to the present invention, X^2 and X^4 defines a sequence of 8-30 amino acids, such as 8-20 amino acids derived from the antigen. This sequence of amino acids derived from an antigen may herein be referred to as an epitope.

Preferably the epitopes used in the scaffold according to the present invention are CTL epitopes. A "CTL inducing peptide" is a HLA Class I binding peptide that is capable of inducing

a CTL response. In other embodiments the epitopes used in the scaffold design according to the present invention are HTL inducing peptides. A "HTL inducing peptide" is a HLA Class II binding peptide that is capable of inducing a HTL response.

5 The term "basic amino acid" as used herein refers to any amino acid including both natural and non-natural amino acids that has an isoelectric point above 6.3 (such as above 7.4) as measured according to Kice & Marvell "Modern Principles of organic Chemistry" (Macmillan, 1974) or Matthews and van Holde "Biochemistry" Cummings Publishing Company, 1996. Included within this definition are Arginine, Lysine, Homoarginine (Har), and Histidine as well as derivatives thereof. Suitable non-natural basic amino acids are e.g. as described in US
10 6,858,396. Suitable positively charged amino acids includes non-natural alpha amino acids available from Bachem AG and includes alpha-amino-glycine, alpha,gamma-diaminobutyric acid, ornithine, alpha, beta-diaminopropionic acid, alpha-difluoromethyl-ornithine, 4-amino-piperidine-4-carboxylic acid, 2,6-diamino-4-hexynoic acid, beta-(1-piperazinyl)-alanine, 4,5-dehydro-lysine, delta-hydroxy-lysine, omega-hydroxy-norarginine, homoarginine, omega-
15 amino-arginine, omega-methyl-arginine, alpha-methyl-histidine, 2,5-diiodo-histidine, 1-methyl-histidine, 3-methyl-histidine, beta-(2-pyridyl)-alanine, beta-(3-pyridyl)-alanine, beta-(2-quinolyl)-alanine, 3-amino-tyrosine, 4-amino-phenylalanine, and spinacine. Furthermore, any mono or dicarboxylic amino acid is a suitable positively charged amino acid.

20 The term "neutral amino acid" as used herein refers to an amino acid that has an isoelectric point above between 4.8 and 6.3 as measured according to Kice & Marvell "Modern Principles of organic Chemistry" (Macmillan, 1974). The term "acidic amino acid" as used herein refers to an amino acid that has an isoelectric point below 4.8 as measured according to Kice & Marvell "Modern Principles of organic Chemistry" (Macmillan, 1974).

25 Unless otherwise indicated amino acids are abbreviated and mentioned by their standard nomenclature known to the person skilled in the art, such as with reference to "nomenclature and symbolism for amino acids and peptides" by the international union of pure and applied chemistry (IUPAC) (www.iupac.org).

30 Citrulline (In this document referred to with the one-letter symbol "B") and/or homocitrulline are well known non-natural amino acids that in some embodiments may be used in the sequence defined by X¹, X³, or X⁵.

In other alternative embodiments, tryptophan or tryptophan derivatives are used in the sequence defined by X¹, X³, or X⁵. Any suitable tryptophan derivatives may be used. As used herein "tryptophan derivatives" means an unnatural modified tryptophan amino acid residue including those disclosed in US 7,232,803, such as tri tert.-butyltryptophan, di-tert-butyl

- tryptophan, 7-benzyloxytryptophan, homotryptophan, 5'-aminoethyltryptophan (available as side chain Boc and N-alpha Fmoc derivative from RSP Amino Acids Analogues Inc, Boston, Mass., USA), N-Acetylhomotryptophan (Toronto Research), 7-Benzyloxytryptophan (Toronto Research), Homotryptophan (Toronto Research), and tryptophan residues which have been substituted at the 1-, 2-, 5- and/or 7-position of the indole ring, positions 1- or 2- being preferred e.g. 5' hydroxy tryptophan.

The term "antibody response" refers to the production of antibodies (e.g., IgM, IgA, IgG) which bind to an antigen of interest, this response is measured for instance by assaying sera by antigen ELISA.

- 10 The term "adjuvant" as used herein refers to any compound which, when delivered together or simultaneously with an antigen, non-specifically enhances the immune response to that antigen. Exemplary adjuvants include but are not limited to oil in water and water in oil adjuvants, aluminum-based adjuvants (e.g., AIOH, AIPO₄, etc), and Montanide ISA 720.
- 15 The terms "patient" and "subject" refer to a mammal that may be treated using the methods of the present invention.

- As used herein, the term "immune response" refers to the reactivity of an organism's immune system in response to an antigen. In vertebrates, this may involve antibody production, induction of cell-mediated immunity, and/or complement activation (e.g., phenomena associated with the vertebrate immune system's prevention and resolution of infection by microorganisms). In preferred embodiments, the term immune response encompasses but is not limited to one or more of a "lymphocyte proliferative response," a "cytokine response," and an "antibody response."

- 25 The term "net charge" as used herein with reference to a peptide sequence refers to the total electric charge of the peptide sequence represented by the sum of charges of each individual amino acid in the peptide sequence, wherein each basic amino acid are given a charge of +1, each acidic amino acid a charge of -1, and each neutral amino acid a charge of 0. Accordingly, the net charge will depend on the number and identities of charged amino acids.

Table 1 – Specific peptides according to the invention

	Reference ID	x1	x2	x3	x4	x5	Placement with

								reference to positions in SEQ ID NO:3; SEQ ID NO:6; SEQ ID NO:7, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:46, and SEQ ID NO:126.	
	P-biotin	R	QIKIWFQN	RR	MKWKK				
	N-Biotin		PVVHRTL	R	QAGDDFSR				
Antigen									
							Modified (m)		
HCV	SP_2	RR	GYIPLVGAPLG	BGR	VARALAHGVRV			X2-seq	x4-seq
HCV	SP_3	R	GYIPLVGAPLG	RR	VARALAHGVRV			135-145	147-157
HCV	SP_4	R	GYIPLVGAPLG	RRR	VARALAHGVRV	R		135-145	147-157
HCV	SP_5	RR	GYIPLVGAPLG	RR	VARALAHGVRV			135-145	147-157
HCV	SP_6	RR	GYIPLVGAPLG	RRR	VARALAHGVRV			135-145	147-157
HCV	SP_7	BR	GYIPLVGAPLG	RR	VARALAHGVRV			135-145	147-157
HCV	SP_8	RRR	GYIPLVGAPLG	BR	VARALAHGVRV			135-145	147-157
HCV	SP_9	R	GYIPLVGAPLG	KKK	VARALAHGVRV			135-145	147-157
HCV	SP_10	R	GYIPLVGAPLG	RRR	VARALAHGVRV			135-145	147-157
HCV	SP_11	KK	GYIPLVGAPLG	KK	VARALAHGVRV			135-145	147-157
HCV	SP_12	W	GYIPLVGAPLG	RR	VARALAHGVRV			135-145	147-157
HCV	SP_13	WW	GYIPLVGAPLG	RR	VARALAHGVRV			135-145	147-157
HCV	SP_14	EE	GYIPLVGAPLG	EE	VARALAHGVRV			135-145	147-157
HCV	SP_15	GG	GYIPLVGAPLG	GG	VARALAHGVRV			135-145	147-157
HCV	SP_16	EE	GYIPLVGAPLG	RR	VARALAHGVRV			135-145	147-157
HCV	SP_17	RR	GYIPLVGAPLG	LRR	VARALAHGVRV			135-145	147-157
HCV	SP21:	WW	GYIPLVGAPLG	RR	VARALAHGVRV			135-145	147-157
HCV	SP22:	WW	GYIPLVGAPLG	RRR	VARALAHGVRV			135-145	147-157
HCV	SP23:	WW	GYIPLVGAPLG	R	VARALAHGVRV			135-145	147-157
HCV	SP24:	R	GYIPLVGAPLG	RR	VARALAHGVRV			135-	147-

								145	157
HCV	51_BIo tin	RR	GYLPAVGAPIG	BR	VIRVIAHGLRL		m	135- 144	147- 157
HCV	51b_BI otin	RR	GYIPLVGAPLG	BR	VARALAHGVRV			135- 145	147- 157
HCV	51_n		GYIPLVGAPLG	G	VARALAHGVRV			135- 145	147- 157
HCV	SP51_1 :	WW	GYLPAVGAPI	RR	VIRVIAHGLRL		m	135- 144	147- 157
HCV	SP1_C *		GYIPLVGAPLG	G	VARALAHGVRV			135- 145	147- 157
HCV	SP2_c	RR	GYIPLVGAPLG	BGR	VARALAHGVRV			135- 145	147- 157
HCV	SP3_c	R	GYIPLVGAPLG	RR	VARALAHGVRV			135- 145	147- 157
HCV	SP4_c	R	GYIPLVGAPLG	RRR	VARALAHGVRV			135- 145	147- 157
HCV	SP5_c	RR	GYIPLVGAPLG	RR	VARALAHGVRV			135- 145	147- 157
HCV	SP6_c	RR	GYIPLVGAPLG	RRR	VARALAHGVRV			135- 145	147- 157
HCV	SP7_c	BR	GYIPLVGAPLG	RR	VARALAHGVRV			135- 145	147- 157
HCV	SP8_c	RRR	GYIPLVGAPLG	BR	VARALAHGVRV			135- 145	147- 157
HCV	SP9_c	R	GYIPLVGAPLG	KKK	VARALAHGVRV			135- 145	147- 157
HCV	SP10_c	R	GYIPLVGAPLG	RRR	VARALAHGVRV			135- 145	147- 157
HCV	SP11_c	KK	GYIPLVGAPLG	KK	VARALAHGVRV			135- 145	147- 157
HCV	SP12_c	W	GYIPLVGAPLG	RR	VARALAHGVRV			135- 145	147- 157
HCV	SP13_c	WW	GYIPLVGAPLG	RR	VARALAHGVRV			135- 145	147- 157
HCV	SP17_c	RR	GYIPLVGAPLG	LRR	VARALAHGVRV			135- 145	147- 157
HCV	SP61_2	RR	NYVTGNIPG	BR	GITFSIFLIVS			163- 171	171- 181
HCV	SP61b_2	WW	NYATGNLPG	RR	CSFSIFLLAL		m	163- 171	171- 181
HCV	SP61_3	WW	NYVTGNIPG	BR	GITFSIFLIVS			163- 171	171- 181
HCV	SP61_4	WW	NYVTGNIPG	RR	GITFSIFLIVS			163- 171	171- 181
HCV	61b_BI otin	RR	NYATGNLPG	RR	GCSFSIFLLAL			163- 171	171- 181
HCV	SP25	RR	VTGNIPGSTYS	GBR	GITFSIYLIVS		m	165- 175	171- 181
HCV	42_BIo tin	RR	IRNLGRVIETLTG	BR	LZGYIPLIGA		m	116- 128	133- 142
HCV	42b_BI otin	RR	SRNLGKVIDTLT C	BR	LMGYIPLVGA			116- 128	133- 142
HCV	42n- BIOTIN		SRNLGKVIDTLT C	GFAD	LMGYIPLVGA			116- 129	133- 142
HCV	SP42_1	WW	IRNLGRVIETLT	RR	LZGYIPLIGA		m	116- 128	133- 142
HCV	SP42b_1	WW	SRNLGKVIDTLT C	RR	LMGYIPLVGA			116- 129	133- 142

HCV	BI310-11_Biotin	RR	GGGQIIGGNYLI P	RB	PBIGVRATB			26-38	42-50
HCV	BI310-11n_Biotin		GGGQIVGGVYLL P	RR	GPRLGVRATR			26-38	42-50
HCV	BI310-11n_sc Biotin	RR	GGGQIVGGVYLL P	RR	GPRLGVRATR			26-38	42-50
HCV	SP11b-1-	WW	GGGQIVGGVYLL P	RR	GPRLGVRAT			26-38	42-50
FLU	BI100-12	BR	LIFLARSALIV		RGSVAHKS			256-266	267-274
FLU	BI100-22b	ED	LIFLARSALIL		RGSVAHKS			255-266	267-274
FLU	120b_B Iotin	BR	LIFLARSALIL	BGR	SALILRGSAVHK			255-266	267-274
FLU	BI100-18b		SAYERMCNIL	KGK	FQTAAQRAMM			217-226	230-239
FLU	BI100-19		SAYERZVNIL	KGK	FQTAAQRAVZ			217-226	230-239
FLU	190_BIotin	BR	TAYERZCNIL	BRGR	FQTVVQBA			217-226	230-237
FLU	190b_B Iotin	BR	IAYERMCNIL	LBRGK	FQTAAQRA			217-226	230-237
FLU	190n-BIOTIN		IAYERMCNIL	KGK	FQTAAQRA			217-226	230-237
FLU	BI100-24b		LFFKCIYRLFKHG L	KR	GPSTEGVPESM			46-59	62-72
FLU	BI100-26	BRR	LFFKTITRLFBHG L	RR	LLSTEGVPNSZ			46-59	62-72
FLU	260_Biotin	BR	GLEPLVIAGILA	RR	GSLVGLLHIVL			23-33	30-40
FLU	260b_Biotin	BR	GSDPLVVAASIV	RR	ASIVGILHLIL			23-33	30-40
CMV	BI 050-sc1	R	NLVPMVATV	RR	NLVPMVATV	B		485-493	485-493
CMV	BI 050-sc2	R	NLVPMVATV	BRR	NLVPMVATV	B		485-493	485-493
CMV	BI 050-sc5	R	NIVPZVVTA	RR	NIVPZVVTA	B	m	485-493	485-493
HIV	N10		PEVIPMFSALS	EGA	TPQDLNNTMLN				
HIV	V10	R	FIIPXFALSG	GRR	ALLYGATPYAIG				
HIV	N13	K	ALGPAATL	EE	MMTACQGVG				
Neg c mod	SP_18	RR	GPVVHLTL	RRR	GQAGDDFS				
Neg c mod	SP_19	RR	GPVVHLTL	RRR	GQAGDDFS				
Neg c	SP_20	RR	GPVVHLTL	RGR	GQAGDDFS				

mod									
HPV		RR	LECVYCKQQLL	RR	EVYDFAFRDLC			35-45	48-58
HPV		RR	GVYDFAFRDLC	RR	GFAFRDLCIVY	R		49-58	52-61
HPV		RR	GVFDYAFRDIN	RR	GFAYRDINLAY	R		49-58	52-61
CMV		RR	GATPVDLLGA	RR	GALNLCLPM	R		498-506	505-514
CMV		RR	GVTPAGLIGV	RR	GALQIBLPL	R		498-506	505-514
HPV		RR	VDIRTLEDLL	RR	GTLGIVCPIG	R		74-83	84-93

As used herein the one-letter-code 'Z' refers to the non-natural amino acid norleucine.

Table 2 – Specific peptides according to the invention

Antigen	X ¹	X ²	X ³	X ⁴	X ⁵
HCV	R	GYIPLVGAPLG	RRR	VARALAHGVRV	R
HCV	R	GYLPAVGAPIG	RRR	VIRVIAHGLRL	R
HCV	RR	GYIPLVGAPLG	RR	VARALAHGVRV	
HCV	RR	GYIPLVGAPLG	RRR	VARALAHGVRV	
HCV	RR	SRNLGKVIDTLTC	RR	LMGYIPLVGA	
HCV	RR	GGGQIVGGVYLLP	RR	GPRLGVRATR	
HCV	W	GYIPLVGAPLG	RR	VARALAHGVRV	
HCV	RR	IRNLGRVIETLTZGYIPLIGA	RR	IRNLGRVIETLTZGYIPLIGA	R
Flu	BR	TAYERZCNIL	BRGR	FQTVVQBA	
cmv	R	NLVPMVATV	BRR	NLVPMVATV	B

As used herein the one-letter-code 'Z' refers to the non-natural amino acid norleucine.

5

Antigens

Examples of viral antigens for use with the present invention include, but are not limited to, e.g., HIV, HCV, CMV, HPV, Influenza, adenoviruses, retroviruses, picornaviruses, etc. Non-limiting example of retroviral antigens such as retroviral antigens from the human

- 10 immunodeficiency virus (HIV) antigens such as gene products of the gag, pol, and env genes, the Nef protein, reverse transcriptase, and other HIV components; hepatitis viral antigens such as the S, M, and L proteins of hepatitis B virus, the pre-S antigen of hepatitis B virus, and other hepatitis, e.g., hepatitis A, B, and C, viral components such as hepatitis C viral RNA; influenza viral antigens such as hemagglutinin and neuraminidase and other influenza
- 15 viral components; measles viral antigens such as the measles virus fusion protein and other measles virus components; rubella viral antigens such as proteins E1 and E2 and other rubella virus components; rotaviral antigens such as VP7sc and other rotaviral components;

cytomegaloviral antigens such as envelope glycoprotein B and other cytomegaloviral antigen components; respiratory syncytial viral antigens such as the RSV fusion protein, the M2 protein and other respiratory syncytial viral antigen components; herpes simplex viral antigens such as immediate early proteins, glycoprotein D, and other herpes simplex viral antigen components; varicella zoster viral antigens such as gpl, gpII, and other varicella zoster viral antigen components; Japanese encephalitis viral antigens such as proteins E, M-E, M-E-NSI, NSI, NS1-NS2A, 80% E, and other Japanese encephalitis viral antigen components; rabies viral antigens such as rabies glycoprotein, rabies nucleoprotein and other rabies viral antigen components. See Fundamental Virology, Second Edition, eds. Fields, B. N. and Knipe, D. M. (Raven Press, New York, 1991) for additional examples of viral antigens.

The epitopes to be incorporated into the scaffold design according to the present invention may be derived from an adenovirus, retrovirus, picornavirus, herpesvirus, rotavirus, hantavirus, coronavirus, togavirus, flavivirus, rhabdovirus, paramyxovirus, orthomyxovirus, bunyavirus, arenavirus, reovirus, papillomavirus, parvovirus, poxvirus, hepadnavirus, or spongiform virus. In certain specific, non-limiting examples, the viral antigen are peptides obtained from at least one of HIV, CMV, hepatitis A, B, and C, influenza, measles, polio, smallpox, rubella; respiratory syncytial, herpes simplex, varicella zoster, Epstein-Barr, Japanese encephalitis, rabies, Influenza, and/or cold viruses.

HCV:

Peptides according to the present invention may comprise a known antigen. For antigens derived from HCV these antigens may be derived from the Core, E1, E2, P7, NS2, NS3, NS4 (NS4A and NS4B) and NS5 (NS5A and NS5B) protein of the Hepatitis C Virus (HCV). The epitopes are those which elicit a HLA class I and/or class II restricted T lymphocyte response in an immunized host. More specific, the HLA class I restricted peptides of the present invention may bind to at least one HLA molecule of the following HLA class I groups: HLA-A*01, HLA-A*02, HLA-A*03, HLA-A*11, HLA-A*24, HLA-B*07, HLA-B*08, HLA-B*35, HLA-B*40, HLA-B*44, HLA-Cw3, HLA-Cw4, HLA-Cw6 or HLA-Cw7. The HLA class II restricted peptides of the present invention may bind to at least one HLA molecule of the following HLA class II groups: HLA-DRB1, -DRB2, -DRB3, -DRB4, -DRB5, -DRB6, -DRB7, -DRB8 or -DRB9.

MHC binding HCV peptides that may be used according to the present invention as epitopes are disclosed in e.g. WO02/34770 (Imperial College Innovations Ltd), WO01/21189 and WO02/20035 (Epimmune), WO04/024182 (Intercell), WO95/25122 (The Scripps Research Institute), WO95/27733 (Government of the USA, Department of Health and Human Services), EP 0935662 (Chiron), WO02/26785 (Immusystems GmbH), WO95/12677 (Innogenetics N.V.), WO97/34621 (Cytel Corp), and EP 1652858 (Innogenetics N.V.).

In other embodiments, the scaffold design according to the present invention comprises a PADRE peptide, such as the universal T cell epitope called PADRE as disclosed in WO95/07707 (Epimmune) the content of which are enclosed herein by reference. A 'PanDR binding peptide or PADRE peptide' is a member of a family of molecules that binds more than
 5 one HLA class II DR molecule. PADRE binds to most HLA-DR molecules and stimulates in vitro and in vivo human helper T lymphocyte (HTL) responses. Alternatively T-help epitopes can be used from universally used vaccines such as tetanos toxoid.

In a further embodiment, the peptides in the composition or polyepitopic peptide are characterized in that they are derived from a HCV protein, or more specifically from at least
 10 one of the following HCV regions selected from the group consisting of Core, E1, E2/NS1, NS2, NS3, NS4A, NS4B, NS5A and NS5B. Even more preferred is that peptides are characterized in that they are present in the HCV consensus sequence of genotype 1a, 1b and/or 3 a.

Other HLA class I and II binding peptides that may be used according to the invention may
 15 be identified by the method as described in WO03/105058 -Algonomics, by the method as described by Epimmune in WO01/21189 and/or by three public database prediction servers, respectively Syfpeithi, BIMAS and nHLAPred. It is also an aspect of this present invention that each peptide may be used within the scaffold design of the invention in combination with the same peptide as multiple repeats, or with any other peptide(s) or epitope(s).

20 CMV:

The epitopes to be incorporated into the scaffold design according to the present invention may be derived from cytomegalovirus (CMV) including CMV glycoproteins gB and gH.

Influenza:

The epitopes to be incorporated into the scaffold design according to the present invention
 25 may be derived from fragments or portions of Influenza hemagglutinin (HA) or Influenza neuraminidase (NA), nucleoprotein (NP), M1, M2, NS1, NEP, PA, PB1, PB1-F2, PB2 for each of the subgroups, such as H1N1, H2N2 og H3N2.

Suitable epitopes may be derived from an HA protein of one, or more than one subtype,
 30 including H1, H2, H3, H4, H5, H6, H7, H8, H9, H10, H11, H12, H13, H14, H15 or H16 or fragment or portion thereof. Examples of subtypes comprising such HA proteins include A/New Caledonia/20/99 (H1N1) A/Indonesia/5/2006 (H5N1), A/chicken/New York/1995, A/herring gull/DE/677/88 (H2N8), A/Texas/32/2003, A/mallard/MN/33/00,

A/duck/Shanghai/1/2000, A/northern pintail/TX/828189/02, A/Turkey/Ontario/6118/68 (H8N4), A/shoveler/Iran/G54/03, A/chicken/Germany/N/1949 (H10N7), A/duck/England/56 (H11N6), A/duck/Alberta/60/76 (H12N5), A/Gull/Maryland/704/77 (H13N6), A/Mallard/Gurjev/263/82, A/duck/Australia/341/83 (H15N8), A/black-headed
 5 gull/Sweden/5/99 (H16N3), B/Lee/40, C/Johannesburg/66, A/PuertoRico/8/34 (H1N1), A/Brisbane/59/2007 (H1N1), A/Solomon Islands 3/2006 (H1N1), A/Brisbane 10/2007 (H3N2), A/Wisconsin/67/2005 (H3N2), B/Malaysia/2506/2004, B/Florida/4/2006, A/Singapore/1/57 (H2N2), A/Anhui/1/2005 (H5N1), A/Vietnam/1194/2004 (H5N1), A/Teal/HongKong/W312/97 (H6N1), A/Equine/Prague/56 (H7N7), A/HongKong/1073/99
 10 (H9N2)).

In some embodiments of the invention, the HA protein may be an H1, H2, H3, H5, H6, H7 or H9 subtype. In other embodiments, the H1 protein may be from the A/New Caledonia/20/99 (H1N1), A/PuertoRico/8/34 (H1N1), A/Brisbane/59/2007 (H1N1), or A/Solomon Islands
 15 3/2006 (H1N1) strain. The H3 protein may also be from the A/Brisbane 10/2007 (H3N2) or A/Wisconsin/67/2005 (H3N2) strain. In other embodiments, the H2 protein may be from the A/Singapore/1/57 (H2N2) strain. The H5 protein may be from the A/Anhui/1/2005 (H5N1), A/Vietnam/1194/2004 (H5N1), or A/Indonesia/5/2005 strain. In other embodiments, the H6 protein may be from the A/Teal/HongKong/W312/97 (H6N1) strain. The H7 protein may be
 20 from the A/Equine/Prague/56 (H7N7) strain. In other embodiments, the H9 protein is from the A/HongKong/1073/99 (H9N2) strain. In other embodiments, the HA protein may be from an influenza virus may be a type B virus, including B/Malaysia/2506/2004 or B/Florida/4/2006. The influenza virus HA protein may be H5 Indonesia.

25 Human immunodeficiency virus (HIV):

For HIV, the epitopes to be incorporated into the scaffold design according to the present invention may be derived from the group consisting of gp120, gp160, gp41, p24gag or p55gag derived from HIV, including members of the various genetic subtypes.

Human papillomavirus (HPV):

30 For HPV, the epitopes to be incorporated into the scaffold design according to the present invention may be derived from the group consisting E1, E2, E3, E4, E6 and E7, L1 and L2 proteins. The epitopes may be derived from any type including types 8, 11, 16, 18, 31, 33, 35, 39, 45, 51, 52, 56, 58, and 59.

Carriers, adjuvants and vehicles - delivery

- The isolated cell-penetrating peptides according to the invention may be delivered by various means and within various compositions, herein referred to as "compositions", "vaccine compositions" or "pharmaceutical compositions". The peptides of the present invention and pharmaceutical and vaccine compositions of the invention are usefull for administration to mammals, particularly humans, to treat and/or prevent virus infection. Vaccine compositions containing the peptides of the invention are administered to a patient infected with the virus in question or to an individual susceptible to, or otherwise at risk for, virus infection to elicit an immune response against the specific antigens and thus enhance the patient's own immune response capabilities.
- Various art-recognized delivery systems may be used to deliver the peptides, into appropriate cells. The peptides can be delivered in a pharmaceutically acceptable carrier or as colloidal suspensions, or as powders, with or without diluents. They can be "naked" or associated with delivery vehicles and delivered using delivery systems known in the art.
- A "pharmaceutically acceptable carrier" or "pharmaceutically acceptable adjuvant" is any suitable excipient, diluent, carrier and/or adjuvant which, by themselves, do not induce the production of antibodies harmful to the individual receiving the composition nor do they elicit protection. Preferably, a pharmaceutically acceptable carrier or adjuvant enhances the immune response elicited by an antigen. Suitable carriers or adjuvant typically comprise one or more of the compounds included in the following non-exhaustive list: large slowly metabolized macromolecules such as proteins, polysaccharides, polylactic acids, polyglycolic acids, polymeric amino acids, amino acid copolymers and inactive virus particles; aluminium hydroxide, aluminium phosphate (see International Patent Application Publication No. WO93/24148), alum ($KAl(SO_4)_2 \cdot 12H_2O$), or one of these in combination with 3-O-deacylated monophosphoryl lipid A (see International Patent Application Publication No. WO93/19780); N-acetyl-muramyl-L-threonyl-D-isoglutamine (see U.S. Patent No. 4,606,918), N-acetyl-normuramyl-L-alanyl-D-isoglutamine, N-acetylmuramyl-L-alanyl-D-isoglutamyl-L-alanine2-(1',2'-dipalmitoyl-sn-glycero-3-hydroxyphosphoryloxy) ethylamine; RIBI (ImmunoChem Research Inc., Hamilton, MT, USA) which contains monophosphoryl lipid A (i.e., a detoxified endotoxin), trehalose-6,6-dimycolate, and cell wall skeleton (MPL + TDM + CWS) in a 2% squalene/Tween 80 emulsion. Any of the three components MPL, TDM or CWS may also be used alone or combined 2 by 2; adjuvants such as Stimulon (Cambridge Bioscience, Worcester, MA, USA), SAF-1 (Syntex); adjuvants such as combinations between QS21 and 3-de-O-acetylated monophosphoryl lipid A (see International Application No. WO94/00153) which may be further supplemented with an oil-in-water emulsion (see, e.g., International Application Nos. WO95/17210, WO97/01640 and WO9856414) in which the oil-in-water emulsion comprises a metabolisable oil and a saponin, or a metabolisable oil, a saponin, and a sterol, or which may be further supplemented with a cytokine (see International Application

- No. WO98/57659); adjuvants such as MF-59 (Chiron), or poly[di(carboxylatophenoxy) phosphazene] based adjuvants (Virus Research Institute); blockcopolymer based adjuvants such as Optivax (Vaxcel, Cytrx) or inulin-based adjuvants, such as Algammulin and Gammalnulin (Anutech); Complete or Incomplete Freund's Adjuvant (CFA or IFA, respectively) or Gerbu preparations (Gerbu Biotechnik); a saponin such as QuilA, a purified saponin such as QS21, QS7 or QS17, -escin or digitonin; immunostimulatory oligonucleotides comprising unmethylated CpG dinucleotides such as [purine-purine-CG-pyrimidine-pyrimidine] oligonucleotides. These immunostimulatory oligonucleotides include CpG class A, B, and C molecules (Coley Pharmaceuticals), ISS (Dynavax), Immunomers (Hybridon).
- 5
- 10 Immunostimulatory oligonucleotides may also be combined with cationic peptides as described, e.g., by Riedl et al. (2002); Immune Stimulating Complexes comprising saponins, for example Quil A (ISCOMS); excipients and diluents, which are inherently non-toxic and non-therapeutic, such as water, saline, glycerol, ethanol, isopropyl alcohol, DMSO, wetting or emulsifying agents, pH buffering substances, preservatives, and the like; a biodegradable
- 15 and/or biocompatible oil such as squalane, squalene, eicosane, tetratetracontane, glycerol, peanut oil, vegetable oil, in a concentration of, e.g., 1 to 10% or 2,5 to 5%; vitamins such as vitamin C (ascorbic acid or its salts or esters), vitamin E (tocopherol), or vitamin A; carotenoids, or natural or synthetic flavanoids; trace elements, such as selenium; any Toll-like receptor ligand as reviewed in Barton and Medzhitov (2002).
- 20 Any of the afore-mentioned adjuvants comprising 3-de-O-acetylated monophosphoryl lipid A, said 3-de-O-acetylated monophosphoryl lipid A may be forming a small particle (see International Application No. WO94/21292).

In any of the aforementioned adjuvants MPL or 3-de-O-acetylated monophosphoryl lipid A can be replaced by a synthetic analogue referred to as RC-529 or by any other amino-alkyl glucosaminide 4-phosphate (Johnson et al. 1999, Persing et al. 2002). Alternatively it can be replaced by other lipid A analogues such as OM-197 (Byl et al. 2003).

25

A "pharmaceutically acceptable vehicle" includes vehicles such as water, saline, physiological salt solutions, glycerol, ethanol, etc. Auxiliary substances such as wetting or emulsifying agents, pH buffering substances, preservatives may be included in such vehicles. Delivery systems known in the art are e.g. lipopeptides, peptide compositions encapsulated in poly-DL-lactide-co-glycolide ("PLG"), microspheres, peptide compositions contained in immune stimulating complexes (ISCOMS), multiple antigen peptide systems (MAPs), viral delivery vectors, particles of viral or synthetic origin, adjuvants, liposomes, lipids, microparticles or microcapsules, gold particles, nanoparticles, polymers, condensing agents, polysaccharides, polyamino acids, dendrimers, saponins, QS21, adsorption enhancing materials, fatty acids or, naked or particle absorbed cDNA.

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Typically, a vaccine or vaccine composition is prepared as an injectable, either as a liquid solution or suspension. Injection may be subcutaneous, intramuscular, intravenous, intraperitoneal, intrathecal, intradermal, or intraepidermal. Other types of administration comprise electroporation, implantation, suppositories, oral ingestion, enteric application, inhalation, aerosolization or nasal spray or drops. Solid forms, suitable for dissolving in, or suspension in, liquid vehicles prior to injection may also be prepared. The preparation may also be emulsified or encapsulated in liposomes for enhancing adjuvant effect.

A liquid formulation may include oils, polymers, vitamins, carbohydrates, amino acids, salts, buffers, albumin, surfactants, or bulking agents. Preferably carbohydrates include sugar or sugar alcohols such as mono-, di-, tri-, oligo- or polysaccharides, or water-soluble glucans. The saccharides or glucans can include fructose, dextrose, lactose, glucose, mannose, sorbose, xylose, maltose, sucrose, dextran, pullulan, dextrin, alpha and beta cyclodextrin, soluble starch, hydroxethyl starch and carboxymethylcellulose, or mixtures thereof. Sucrose is most preferred. "Sugar alcohol" is defined as a C4 to C8 hydrocarbon having an -OH group and includes galactitol, inositol, mannitol, xylitol, sorbitol, glycerol, and arabitol. Mannitol is most preferred. These sugars or sugar alcohols mentioned above may be used individually or in combination. There is no fixed limit to the amount used as long as the sugar or sugar alcohol is soluble in the aqueous preparation. Preferably, the sugar or sugar alcohol concentration is between 1,0 % (w/v) and 7,0 % (w/v), more preferable between 2,0 and 6,0 % (w/v). Preferably amino acids include levorotary (L) forms of carnitine, arginine, and betaine; however, other amino acids may be added. Preferred polymers include polyvinylpyrrolidone (PVP) with an average molecular weight between 2,000 and 3,000, or polyethylene glycol (PEG) with an average molecular weight between 3,000 and 5,000. It is also preferred to use a buffer in the composition to minimize pH changes in the solution before lyophilization or after reconstitution. Any physiological buffer may be used, but citrate, phosphate, succinate, and glutamate buffers or mixtures thereof are preferred. Most preferred is a citrate buffer. Preferably, the concentration is from 0,01 to 0,3 molar. Surfactants that can be added to the formulation are shown in EP patent applications No. EP 0 270 799 and EP 0 268 110.

Additionally, polypeptides can be chemically modified by covalent conjugation to a polymer to increase their circulating half-life, for example. Preferred polymers, and methods to attach them to peptides, are shown in U.S. Patent Nos. 4,766,106; 4,179,337; 4,495,285; and 4,609,546. Preferred polymers are polyoxyethylated polyols and polyethylene glycol (PEG). PEG is soluble in water at room temperature and has the general formula:

$R(O-CH_2-CH_2)_nO-R$ where R can be hydrogen, or a protective group such as an alkyl or alkanol group. Preferably, the protective group has between 1 and 8 carbons, more

preferably it is methyl. The symbol n is a positive integer, preferably between 1 and 1.000, more preferably between 2 and 500. The PEG has a preferred average molecular weight between 1000 and 40.000, more preferably between 2000 and 20.000, most preferably between 3.000 and 12.000. Preferably, PEG has at least one hydroxy group, more preferably it is a terminal hydroxy group. It is this hydroxy group which is preferably activated.
However, it will be understood that the type and amount of the reactive groups may be varied to achieve a covalently conjugated PEG/polypeptide of the present invention.

Water soluble polyoxyethylated polyols are also useful in the present invention. They include polyoxyethylated sorbitol, polyoxyethylated glucose, polyoxyethylated glycerol (POG), etc.
POG is preferred. One reason is because the glycerol backbone of polyoxyethylated glycerol is the same backbone occurring naturally in, for example, animals and humans in mono-, di-, triglycerides. Therefore, this branching would not necessarily be seen as a foreign agent in the body. The POG has a preferred molecular weight in the same range as PEG. The structure for POG is shown in Knauf et al., 1988, and a discussion of POG/IL-2 conjugates is found in U.S. Patent No. 4,766,106.

Another drug delivery system for increasing circulatory half-life is the liposome. The peptides and nucleic acids of the invention may also be administered via liposomes, which serve to target a particular tissue, such as lymphoid tissue, or to target selectively infected cells, as well as to increase the half-life of the peptide and nucleic acids composition. Liposomes include emulsions, foams, micelles, insoluble monolayers, liquid crystals, phospholipid dispersions, lamellar layers and the like. In these preparations, the peptide or nucleic acids to be delivered is incorporated as part of a liposome or embedded, alone or in conjunction with a molecule which binds to a receptor prevalent among lymphoid cells, such as monoclonal antibodies which bind to the CD45 antigen, or with other therapeutic or immunogenic compositions. Thus, liposomes either filled or decorated with a desired peptide or nucleic acids of the invention can be directed to the site of lymphoid cells, where the liposomes then deliver the peptide and nucleic acids compositions. Liposomes for use in accordance with the invention are formed from standard vesicle-forming lipids, which generally include neutral and negatively charged phospholipids and a sterol, such as cholesterol. The selection of lipids is generally guided by consideration of, e.g., liposome size, acid lability and stability of the liposomes in the blood stream. A variety of methods are available for preparing liposomes, as described in, e.g., Szoka et al, 1980, and U.S. Patent Nos. 4,235,871, 4,501,728, 4,837,028, and 5,019,369.

For targeting cells of the immune system, a ligand to be incorporated into the liposome can include, e.g., antibodies or fragments thereof specific for cell surface determinants of the desired immune system cells. A liposome suspension containing a peptide may be

administered intravenously, locally, topically, etc. in a dose which varies according to, inter alia, the manner of administration, the peptide being delivered, and the stage of the disease being treated. For example, liposomes carrying either immunogenic polypeptides are known to elicit CTL responses in vivo (Reddy et al., 1992; Collins et al., 1992; Fries et al., 1992; Nabel et al., 1992).

After the liquid pharmaceutical composition is prepared, it is preferably lyophilized to prevent degradation and to preserve sterility. Methods for lyophilizing liquid compositions are known to those of ordinary skill in the art. Just prior to use, the composition may be reconstituted with a sterile diluent (Ringer's solution, distilled water, or sterile saline, for example) which may include additional ingredients. Upon reconstitution, the composition is preferably administered to subjects using those methods that are known to those skilled in the art.

Use of the peptides for evaluating immune responses.

The peptides according to the present invention may be used as diagnostic reagents. For example, a peptide of the invention may be used to determine the susceptibility of a particular individual to a treatment regimen which employs the peptide or related peptides, and thus may be helpful in modifying an existing treatment protocol or in determining a prognosis for an affected individual. In addition, the peptides may also be used to predict which individuals will be at substantial risk for developing a chronic virus infection.

Accordingly, the present invention relates to a method of determining the outcome for a subject exposed to a virus, comprising the steps of determining whether the subject has an immune response to one or more peptides according to the present invention.

In a preferred embodiment of the invention, the peptides as described herein can be used as reagents to evaluate an immune response. The immune response to be evaluated can be induced by using as an immunogen any agent that may result in the production of antigen-specific CTLs or HTLs that recognize and bind to the peptide(s) to be employed as the reagent. The peptide reagent need not be used as the immunogen. Assay systems that can be used for such an analysis include relatively recent technical developments such as tetramers, staining for intracellular lymphokines and interferon release assays, or ELISPOT assays.

For example, a peptide of the invention may be used in a tetramer staining assay to assess peripheral blood mononuclear cells for the presence of antigen-specific CTLs following exposure to an antigen or an immunogen. The HLA- tetrameric complex is used to directly visualize antigen-specific CTLs (see, e.g., Ogg et al., 1998; and Altman et al., 1996) and

determine the frequency of the antigen-specific CTL population in a sample of peripheral blood mononuclear cells. A tetramer reagent using a peptide of the invention may be generated as follows: a peptide that binds to an HLA molecule is refolded in the presence of the corresponding HLA heavy chain and beta2-microglobulin to generate a trimolecular
5 complex. The complex is biotinylated at the carboxyl terminal end of the heavy chain at a site that was previously engineered into the protein. Tetramer formation is then induced by the addition of streptavidin. By means of fluorescently labeled streptavidin, the tetramer can be used to stain antigen-specific cells. The cells may then be identified, for example, by flow cytometry. Such an analysis may be used for diagnostic or prognostic purposes. Cells
10 identified by the procedure can also be used for therapeutic purposes. As an alternative to tetramers also pentamers or dimers can be used (Current Protocols in Immunology (2000) unit 17.2 supplement 35)

Peptides of the invention may also be used as reagents to evaluate immune recall responses. (see, e.g., Bertoni et al., 1997 and Perma et al., 1991.). For example, patient PBMC samples
15 from individuals with HCV infection may be analyzed for the presence of antigen-specific CTLs or HTLs using specific peptides. A blood sample containing mononuclear cells may be evaluated by cultivating the PBMCs and stimulating the cells with a peptide of the invention. After an appropriate cultivation period, the expanded cell population may be analyzed, for example, for cytotoxic activity (CTL) or for HTL activity.

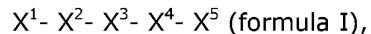
20 The peptides may also be used as reagents to evaluate the efficacy of a vaccine.

PBMCs obtained from a patient vaccinated with an immunogen may be analyzed using, for example, either of the methods described above. The patient is HLA typed, and peptide epitope reagents that recognize the allele-specific molecules present in that patient are selected for the analysis. The immunogenicity of the vaccine is indicated by the presence of
25 epitope-specific CTLs and/or HTLs in the PBMC sample.

The peptides of the invention may also be used to make antibodies, using techniques well known in the art (see, e.g. CURRENT PROTOCOLS IN IMMUNOLOGY, Wiley/Greene, NY; and Antibodies A Laboratory Manual, Harlow and Lane, Cold Spring Harbor Laboratory Press, 1989). Such antibodies include those that recognize a peptide in the context of an HLA
30 molecule, i.e., antibodies that bind to a peptide-MHC complex.

Specific embodiments of the invention

The present invention concerns a peptide an isolated cell-penetrating peptide comprising the following structure



- 5 wherein X^1 and X^3 independently defines a linear sequence of any 1, 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof; X^2 defines a linear sequence of 8-30 amino acids derived from an antigen; X^4 defines a linear sequence of 8-30 amino acids derived from said antigen, said sequence X^4 being different from X^2 ; and wherein X^5 is any one optional amino acid selected from a basic amino
10 acid, citrulline, tryptophan, or a derivative thereof.

In some specific embodiments the sequence of X^4 is identical to the sequence of X^2 .

In some embodiments the isolated cell-penetrating peptide has a total of not more than 60 amino acids.

- 15 In some embodiments X^1 defines a linear sequence of any 1, 2, 3, or 4 amino acids independently selected from any basic amino acid, tryptophan, or a derivative thereof.

In some embodiments X^1 defines a linear sequence in any order of one citrulline and any 1, 2, or 3 amino acids independently selected from any basic amino acid, tryptophan, or a derivative thereof.

- 20 In some embodiments X^3 defines a linear sequence of any 1, 2, 3, or 4 amino acids independently selected from any basic amino acid.

In some embodiments X^1 defines a linear sequence of any 1, 2, 3, or 4 amino acids independently selected from any basic amino acid.

In some embodiments X^3 defines a linear sequence in any order of one citrulline and of any 1, 2, or 3 amino acids independently selected from any basic amino acid.

- 25 In some embodiments the sequence of amino acids defined by $X^2-X^3-X^4$ of formula I as defined in claim 1 is not found in the native sequence of said antigen.

In some embodiments the sequence of amino acids defined by X^1 - X^2 - X^3 - X^4 of formula I as defined in claim 1 is not found in the native sequence of said antigen.

In some embodiments the sequence of amino acids defined by X^1 - X^2 - X^3 - X^4 - X^5 of formula I as defined in claim 1 is not found in the native sequence of said antigen.

- 5 In some embodiments the peptide is demonstrated to translocate across a plasma membrane in the assay based on biotinylation of peptides as described in example 1.

In some embodiments the 1, 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof, is a basic amino acid.

- 10 In some embodiments the 1, 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof, is tryptophan, or a derivative thereof.

In some embodiments the 1, 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof, is citrulline, or a derivative thereof.

- 15 In some embodiments the one optional amino acid selected from a basic amino acid, citrulline, tryptophan, or a derivative thereof is selected from Arg, Lys, and His.B).

In some embodiments the one optional amino acid selected from a basic amino acid, citrulline, tryptophan or a derivative thereof is tryptophan, or a derivative thereof.

In some embodiments the one optional amino acid selected from a basic amino acid, citrulline, tryptophan or a derivative thereof is citrulline, or a derivative thereof.

- 20 In some embodiments X^2 and/or X^4 defines a sequence identical to the native sequence of said antigen.

In some embodiments the peptide is capable of inducing a T-lymphocyte response.

In some embodiments the peptide is capable of inducing a CD4+ and/or a CD8+ T-lymphocyte response.

- 25 In some embodiments the antigen is a viral protein, such as a capsid protein.

In some embodiments the viral protein is selected from a protein of the Hepatitis C virus, such as a core protein; protein of influenza virus, such as an M2 protein.

In some embodiments the viral protein of Hepatitis C virus is selected from HCV consensus sequence of genotype 1, such as subtypes 1a and 1b, genotype 2 such as 2a and 2b and
5 genotype 3, such as 3a.

In some embodiments the cell-penetrating peptide is of 19-60 amino acids, such as of 20-60 amino acids, such as of 21-60 amino acids, such as of 22-60 amino acids, such as of 23-60 amino acids, such as of 24-60 amino acids, such as of 25-60 amino acids, such as of 26-60 amino acids, such as of 27-60 amino acids, such as of 28-60 amino acids, such as of 29-60
10 amino acids, such as of 30-60 amino acids, such as of 31-60 amino acids, such as of 32-60 amino acids, such as of 33-60 amino acids, such as of 34-60 amino acids, such as of 35-60 amino acids.

In some embodiments the cell-penetrating peptide is of 18-60 amino acids, such as 18-59 amino acids, such as 18-58 amino acids, such as 18-57 amino acids, such as 18-56 amino
15 acids, such as 18-55 amino acids, such as 18-54 amino acids, such as 18-53 amino acids, such as 18-52 amino acids, such as 18-51 amino acids, such as 18-50 amino acids, such as 18-49 amino acids, such as 18-48 amino acids, such as 18-47 amino acids, such as 18-46 amino acids, such as 18-45 amino acids, such as 18-44 amino acids, such as 18-43 amino acids, such as 18-42 amino acids, such as 18-41 amino acids, such as 18-40 amino acids,
20 such as 18-39 amino acids, such as 18-38 amino acids, such as 18-37 amino acids, such as 18-35 amino acids, such as of 18-34 amino acids, such as of 18-33 amino acids, such as of 18-32 amino acids, such as of 18-31 amino acids, such as of 18-30 amino acids, such as of 18-29 amino acids, such as of 18-28 amino acids, such as of 18-27 amino acids, such as of 18-26 amino acids, such as of 18-25 amino acids, such as of 18-24 amino acids, such as of
25 18-23 amino acids, such as of 18-22 amino acids, such as of 18-21 amino acids, such as of 18-20 amino acids, such as of 18-19 amino acids.

In some embodiments the net charge of X^2 is below or equal to 0.

In some embodiments the net charge of X^2 is below or equal to 0; and X^1 and X^3 defines a sequence of 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline,
30 tryptophan, or a derivative thereof.

In some embodiments the net charge of X^2 is below or equal to 0; and the net charge of X^4 is above or equal to 1.

In some embodiments the net charge of X^2 is below or equal to 0; the net charge of X^4 is above or equal to 1; and X^1 and X^3 defines a sequence of 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof.

In some embodiments the net charge of X^2 and X^4 are below or equal to 0.

- 5 In some embodiments the net charge of X^2 and X^4 are below or equal to 0; and X^1 and X^3 defines a sequence of 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof.

In some embodiments the net charge of X^2 and X^4 are above or equal to 1.

- 10 In some embodiments the net charge of X^2 and X^4 are above or equal to 1 and X^1 and X^3 defines a sequence of 1 or 2 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof.

In some embodiments the net charge of X^2 is above or equal to 1; and the net charge of X^4 is below or equal to 0.

- 15 In some embodiments the net charge of X^2 is above or equal to 1; the net charge of X^4 is below or equal to 0; X^1 defines a sequence of 1 or 2 amino acids with a positive charge; and X^3 defines a sequence of 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof.

In some embodiments the basic amino acid is independently selected from Arg, Lys, and His.

- 20 In some embodiments X^2 and/or X^4 comprises the sequence of amino acids defined by position 135-157 of SEQ ID NO:3, or a fragment or variant thereof.

- 25 In some embodiments X^2 and/or X^4 consist of a sequence selected from GYIPLVGAPLG, GYLPAVGAPIG, GYLPAVGAPI, NYVTGNIPG, NYATGNLPG, NYATGNLPG, VTGNIPGSTYS, IRNLGRVIETLTG, SRNLGKVIDTLTC, IRNLGRVIETLT, GGGQIIGGNYLIP, GGGQIVGGVYLLP, LIFLARSALIV, LIFLARSALIL, LIFLARSALIL, SAYERM CNIL, SAYERZVNIL, TAYERZCNIL, IAYERMCNIL, IAYERMCNIL, LFFKCIYRLFKHGL, LFFKTITRLFBHGL, GLEPLVIAGILA, GSDPLVVAASIV, NLVPMVATV, NLVPMVATV, NIVPZVVTA, PEVIPMFSALS, FIIPXFALSG, ALGPAATL, GPVVHRTL, LECVYCKQQLL, GYDFAFRDLC, GVFDYAFRDIN, GATPVDLLGA, GVTPAGLIGV, VARALAHGVRV, VIRVIAHGLRL, GITFSIFLIVS, CSFSIFLLAL, GCSFSIFLLAL, GITFSIYLIVS, LZGYIPLIGA, LMGYIPLVGA, LZGYIPLIGA, PBIGVRATB, GPRLGVRATR,

GPRLGVRAT, RGSVAHKS, SALILRGSAHK, FQTAAQRAMM, FQTAAQRAVZ, FQTVVQBA, FQTAAQRA, GPSTEGVPESM, LLSTEGVPNSZ, GSLVGLLHIVL, ASIVGILHLIL, NLVPMVATV, NIVPZVVTA, TPQDLNTMLN, ALLYGATPYAIG, MMTACQGVG, GQAGDDFS, EVYDFAFRDLC, GFAFRDLCIVY, GFAYRDINLAY, GALNLCLPM, and GALQIBLPL, IRNLGRVIETLTZGYIPLIGA, or a
 5 fragment or variant thereof.

In some embodiments X^2 and/or X^4 consist of a sequence derived from an amino acid sequence selected from SEQ ID NO:1, SEQ ID NO:2, SEQ ID NO:3, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:6, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:10, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, SEQ ID NO:14, SEQ ID NO:15, SEQ ID NO:16, SEQ
 10 ID NO:17, SEQ ID NO:18, SEQ ID NO:19, SEQ ID NO:20, SEQ ID NO:21, SEQ ID NO:22, SEQ ID NO:23, SEQ ID NO:24, SEQ ID NO:25, SEQ ID NO:26, SEQ ID NO:27, SEQ ID NO:28, SEQ ID NO:29, SEQ ID NO:30, SEQ ID NO:31, SEQ ID NO:32, SEQ ID NO:33, SEQ ID NO:34, SEQ ID NO:35, SEQ ID NO:36, SEQ ID NO:37, SEQ ID NO:38, SEQ ID NO:39, SEQ ID NO:40, SEQ ID NO:41, SEQ ID NO:42, SEQ ID NO:43, SEQ ID NO:44, SEQ ID
 15 NO:45, SEQ ID NO:46, SEQ ID NO:126, and SEQ ID NO:198, or a fragment or variant thereof.

In some embodiments the peptide according to the invention is selected from RRGYIPLVGAPLGBGRVARALAHGVRV, RGYIPLVGAPLGRRVARALAHGVRV, RGYIPLVGAPLGRRRVARALAHGVRV, RRGYIPLVGAPLGRRVARALAHGVRV,
 20 RRGYIPLVGAPLGRRRVARALAHGVRV, BRGYIPLVGAPLGRRVARALAHGVRV, RRRGYIPLVGAPLGBRVARALAHGVRV, RGYIPLVGAPLGKKKVARALAHGVRV, RGYIPLVGAPLGRRRVARALAHGVRV, KKGYIPLVGAPLGKKVARALAHGVRV, WGYIPLVGAPLGRRVARALAHGVRV, WWGYIPLVGAPLGRRVARALAHGVRV, EEGYIPLVGAPLGEEVARALAHGVRV, GGGYIPLVGAPLGGGVARALAHGVRV,
 25 EEGYIPLVGAPLGRRVARALAHGVRV, RRGYIPLVGAPLGRRVARALAHGVRV, WWGYIPLVGAPLGRRVARALAHGVRV, WWGYIPLVGAPLGRRRVARALAHGVRV, WWGYIPLVGAPLGRVARALAHGVRV, RGYIPLVGAPLGRRVARALAHGVRV, RRGYIPLVGAPLGBRVARALAHGVRV, GYIPLVGAPLGGVARALAHGVRV, WWGYIPLVGAPLRRVIRVIAHGLRL,
 30 GYIPLVGAPLGGVARALAHGVRV, RRGYIPLVGAPLGBGRVARALAHGVRV, RGYIPLVGAPLGRRVARALAHGVRV, RGYIPLVGAPLGRRRVARALAHGVRV, RRGYIPLVGAPLGRRRVARALAHGVRV, BRGYIPLVGAPLGRRVARALAHGVRV, RRRGYIPLVGAPLGBRVARALAHGVRV, RGYIPLVGAPLGKKKVARALAHGVRV, RGYIPLVGAPLGRRRVARALAHGVRV,
 35 KKGYIPLVGAPLGKKVARALAHGVRV, WGYIPLVGAPLGRRVARALAHGVRV, WWGYIPLVGAPLGRRVARALAHGVRV, RRGYIPLVGAPLGRRVARALAHGVRV, RRNYVTGNIPGBRGITFSIFLIVS, WWNYATGNLPGRRCSFSIFLLAL,

WWNYVTGNIPGBRGITFSIFLIVS, WWNYVTGNIPGRRGITFSIFLIVS,
 RRNYATGNLPGRRGCSFSIFLLAL, RRVGTGNIPGSTYSGBRGITFSIYLIVS,
 RRIRNLGRVIETLTGBRLZGYIPLIGA, RRSRNLGKVIDTLTCBRLMGYIPLVGA,
 SRNLGKVIDTLTCGFADLMGYIPLVGA, WWIRNLGRVIETLTRRLZGYIPLIGA,
 5 WWSRNLGKVIDTLTCRRLMGYIPLVGA, RRGGGQIIGGNYLIPRBPBIGVRATB,
 GGGQIVGGVYLLPRRGPRLGVRATR, RRGGGQIVGGVYLLPRRGPRLGVRATR,
 WWGGGQIVGGVYLLPRRGPRLGVRAT, , BRLIFLARSALIVRGSVAHKS,
 EDLIFLARSALILRGSVAHKS, BRLIFLARSALILBGRSALILRGSVAHK,
 SAYERMCNILKGKFQTAAQRAMM, SAYERZVNILKGKFQTAAQRAVZ,
 10 BRTAYERZCNILBRGRFQTVVQBA, BRIAYERMCNILLBRGKFQTAAQRA,
 IAYERMCNILKGKFQTAAQRA, LFFKCIYRLFKHGLKRGPESTEGVPESM,
 BRRLFFKTITRLFBHGLRRLSTEGVPNSZ, BRGLEPLVIAGILARRGSLVGLLHIVL,
 BRGSDPLVVAASIVRRASIVGILHLIL, , RNLVPMVATVRRNLVPMVATVB,
 RNLVPMVATVBRRNLVPMVATVB, RNIVPZVVTARRNIVPZVVTAB, ,
 15 PEVIPMFSALEGATPQDLNMTLN, RFIIPXFTALSGGRRALLYGATPYAIG,
 KALGPAATLEEMMTACQGVG, , RRGPVVHLTLRRRGQAGDDFS, RRGPVVHLTLRRRGQAGDDFS,
 RRGPVVHLTLRGRRGQAGDDFS, RRLECVYCKQQLLREVYDFAFRDLC,
 RRGVYDFAFRDLCRRGFAFRDLCIVYR, RRGVFDYAFRDINRRGFAYRDINLAYR,
 RRGATPVDLLGARRGALNLCLPMR, RRGVTPAGLIGVRRGALQIBLPLR,
 20 RGYLPAVGAPIGRRRVIRVIAHGLRLR, RRSRNLGKVIDTLTCRRLMGYIPLVGA,
 RRIRNLGRVIETLTZGYIPLIGARRIRNLGRVIETLTZGYIPLIGAR, or a fragment or variant
 thereof.

In some embodiments the peptide consist of a sequence selected from X^1 -NYVTGNIPG- X^3 -
 GITFSIYLIVS; X^1 -IRNLGRVIETLT- X^3 -LZGYIPLIGA; X^1 -GYLPAVGAPI- X^3 -VIRVIAHGLRL; X^1 -
 25 GGGQIIGGNYLIP- X^3 -PBIGVRATB; X^1 -NYATGNLPG- X^3 -GCSFSIFLLAL; X^1 -SRNLGKVIDTLTC- X^3 -
 LMGYIPLVGA; X^1 -GYIPLVGAPL- X^3 -VARALAHGVRV; X^1 -GGGQIVGGVYLLP- X^3 -PRLGVRATR; X^1 -
 LTFLVRSVLLI- X^3 -GSVLIVRGSVH; X^1 -TAYERZCNIL- X^3 -GRFQTVVQBA; X^1 -SDPLVVAASIV- X^3 -
 ASIVGILHLIL; X^1 -LIFLARSALIL- X^3 -SALILRGSVAH; X^1 -IAYERMCNIL- X^3 -GKFQTAAQRA; and X^1 -
 LEPLVIAGILA- X^3 -GSLVGLLHIVL; X^1 -NLVPMVATV- X^3 -NLVPMATV; X^1 -GYLPAVGAPIG- X^3 -
 30 VIRVIAHGLRL; X^1 -IRNLGRVIETLTG- X^3 -LZGYIPLIGA; X^1 -GVYDFAFRDLC- X^3 -GFAFRDLCIVYR, X^1 -
 GVFYDFAFRDIN- X^3 -GFAYRDINLAYR, X^1 -GATPVDLLGA- X^3 -GALNLCLPMR, X^1 -GVTPAGLIGV- X^3 -
 GALQIBLPLR, and X^1 -IRNLGRVIETLTZGYIPLIGA- X^3 -IRNLGRVIETLTZGYIPLIGA; optionally
 with an X^5 in the C-terminal of the peptide wherein X^1 , X^3 and X^5 refers to X^1 , X^3 , and X^5 of
 formula I.

35 In some embodiments the peptide comprises one or more cysteine.

In some embodiments the peptide contain intramolecular bonds, such as intramolecular disulfide (S-S) bonds between two cys residues.

In other embodiments the peptide contains intramolecular bonds, such as in the form of a acylal moiety (COO-CH₂-OOC, COO-CHR-OOC or COO-CR₂-OOC).

- 5 In some embodiments the N- and/or C-terminal amino acid in X² is a hydrophilic or polar amino acid.

In some embodiments the N-terminal amino acid in X⁴ is a hydrophilic or polar amino acid.

- 10 In some embodiments the peptide according to the invention is not more than 58 amino acids, such as not more than 56, 54, 52, 50, 48, 46, 44, 42, 40, 38, 36, 34, 32, 30, 28, 26, 24, 22, 20, 18 amino acid residues.

In some embodiments X¹ consist of 2 or 3 amino acids.

In some embodiments X¹ consist of WW, BR, or RR.

In some embodiments X¹ consist of R, or W.

In some embodiments X³ consist of 2 or 3 amino acids.

- 15 In some embodiments X³ consist of WW, BR, or RR.

In some embodiments X³ consist of RRR, BRR, or BRGR.

In some embodiments an isolated peptide according to the invention consists of a sequence of X² or X⁴ as defined in table 1 or in table 2.

- 20 In some embodiments an isolated peptide according to the invention comprises a sequence of X² and/or X⁴ as defined in table 1 or in table 2, or a fragment thereof.

In some embodiments X² defines a sequence of 8-25 amino acids, such as 8-20 amino acids, such as 8-15 amino acids, such as 8-11 amino acids.

In some embodiments X² defines a sequence of less than 25, such as less than 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7 or 6 amino acids.

In some embodiments X^2 defines a sequence of more than 8, such as more than 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29 amino acids.

In some embodiments X^4 defines a sequence of 8-25 amino acids, such as 8-20 amino acids, such as 8-15 amino acids, such as 8-11 amino acids.

- 5 In some embodiments X^4 defines a sequence of less than 25, such as less than 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7 or 6 amino acids.

In some embodiments X^4 defines a sequence of more than 8, such as more than 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29 amino acids.

- 10 In some embodiments the dimer peptide according to the invention consist of two identical peptide monomers.

In some embodiments the immunogenic composition according to the invention is in the form of a vaccine composition.

- 15 In some embodiments, the cell-penetrating peptide of the invention comprises at most 60, at most 59, at most 58, at most 57, at most 56, at most 55, at most 54, at most 53, at most 52, at most 51, at most 50, at most 49, at most 48, at most 47, at most 46, at most 45, at most 44, at most 43, at most 42, at most 41, at most 40, at most 39, at most 38, at most 37, at most 36, at most 35, at most 34, at most 33, at most 32, at most 31, at most 30, at most 29, at most 28, at most 27, at most 26, at most 25, at most 24, at most 23, at most 22, at most 21, at most 20, at most 19, at most 18 amino acids.

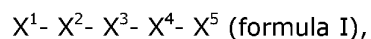
- 20 In some embodiments, the cell-penetrating peptide of the invention comprises at least 18, at least 19, at least 20, at least 21, at least 22, at least 23, at least 24, at least 25, at least 26, at least 27, at least 28, at least 29, at least 30, at least 31, at least 32, at least 33, at least 34, at least 35 at least 36, at least 37, at least 38, at least 39, at least 40, at least 41, at least 42, at least 43, at least 44, at least 45, at least 46, at least 47, at least 48, at least 49,
25 at least 50, at least 51, at least 52, at least 53, at least 54, at least 55, at least 56, at least 57, at least 58, at least 59, at least 60 amino acid residues.

- In some embodiments, the cell-penetrating peptide of the invention consists of 18 amino acid residues or 19 amino acid residues or 20 amino acid residues or 21 amino acid residues or 22 amino acid residues or 23 amino acid residues or 24 amino acid residues or 25 amino acid
30 residues or 26 amino acid residues or 27 amino acid residues or 28 amino acid residues or 29 amino acid residues or 30 amino acid residues or 31 amino acid residues or 32 amino acid

- residues or 33 amino acid residues or 34 amino acid residues or 35 amino acid residues or 36 amino acid residues or 37 amino acid residues or 38 amino acid residues or 39 amino acid residues or 40 amino acid residues or 41 amino acid residues or 42 amino acid residues or 43 amino acid residues or 44 amino acid residues or 45 amino acid residues or 46 amino acid residues or 47 amino acid residues or 48 amino acid residues or 49 amino acid residues or 50 amino acid residues or 51 amino acid residues or 52 amino acid residues or 53 amino acid residues or 54 amino acid residues or 55 amino acid residues or 56 amino acid residues or 57 amino acid residues or 58 amino acid residues or 59 amino acid residues or 60 amino acid residues.
- 5
- 10 In some embodiments the cell-penetrating peptide of the invention does not consist of the following sequence RFIIP[Nle]FTALSGGRRALLYGATPYAIG, where Nle denotes a nor-leucine.
- In some embodiments X^2 and/or X^4 is not derived from HIV.
- In some embodiments X^4 is a linear sequence of less than 12 amino acids.
- In some embodiments X^2 is a linear sequence of less than 12 amino acids.
- 15 In some embodiments X^2 and/or X^4 do not contain nor-leucine.
- In some embodiments X^2 do not contain nor-leucine.
- In some embodiments X^2 and/or X^4 only contains natural amino acids.
- In some embodiments X^2 only contains natural amino acids.
- In some embodiments X^2 only contains natural amino acids if derived from HIV.
- 20 In some embodiments X^2 and/or X^4 is derived from HCV, CMV, HPV, Influenza, adenoviruses, or picornaviruses.

Numbered embodiments according to the invention:

1. An isolated cell-penetrating peptide comprising the following structure



- wherein X^1 and X^3 independently defines a linear sequence of any 1, 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof; X^2 defines a linear sequence of 8-30 amino acids derived from an antigen; X^4 defines a linear sequence of 8-30 amino acids derived from said antigen, said sequence X^4 being different from X^2 ; and wherein X^5 is any one optional amino acid selected from a basic amino acid, citrulline, tryptophan, or a derivative thereof.
- 5
2. The isolated peptide according to embodiment 1, wherein X^1 defines a linear sequence of any 1, 2, 3, or 4 amino acids independently selected from any basic amino acid, tryptophan, or a derivative thereof.
- 10
3. The isolated peptide according to embodiment 1, wherein X^1 defines a linear sequence in any order of one citrulline and any 1, 2, or 3 amino acids independently selected from any basic amino acid, tryptophan, or a derivative thereof.
4. The isolated peptide according to any one of embodiment 1-3, wherein X^3 defines a linear sequence of any 1, 2, 3, or 4 amino acids independently selected from any basic amino acid.
- 15
5. The isolated peptide according to any one of embodiment 1-3, wherein X^3 defines a linear sequence in any order of one citrulline and of any 1, 2, or 3 amino acids independently selected from any basic amino acid.
6. The isolated peptide according to any one of embodiment 1-5, wherein the sequence of amino acids defined by X^2 - X^3 - X^4 of formula I as defined in embodiment 1 is not found in the native sequence of said antigen.
- 20
7. The isolated peptide according to any one of embodiment 1-6, wherein said peptide is demonstrated to translocate across a plasma membrane in the assay based on biotinylation of peptides as described in example 1.
- 25
8. The isolated peptide according to any one of embodiments 1-7, wherein said one optional amino acid selected from a basic amino acid, citrulline, tryptophan, or a derivative thereof is selected from Arg, Lys, and His.
9. The isolated peptide according to any one of embodiments 1-8, wherein X^2 and/or X^4 defines a sequence identical to the native sequence of said antigen.

10. The isolated peptide according to any one of embodiments 1-9, wherein said peptide is capable of inducing a T lymphocyte response.
11. The isolated peptide according to any one of embodiments 1-10, wherein said antigen is a viral protein, such as a capsid protein.
- 5 12. The isolated peptide according to embodiment 11, wherein said viral protein is selected from a protein of the Hepatitis C virus, such as a core protein; protein of influenza virus, such as an M2 protein.
13. The isolated peptide according to embodiment 11, wherein said viral protein of Hepatitis C virus is selected from HCV consensus sequence of genotype 1, such as subtypes
10 1a and 1b, genotype 2 such as 2a and 2b and genotype 3, such as 3a.
14. The isolated peptide according to any one of embodiments 1-13, which cell-penetrating peptide is of 19-60 amino acids, such as of 20-60 amino acids, such as of 21-60 amino acids, such as of 22-60 amino acids, such as of 23-60 amino acids, such as of 24-60 amino acids, such as of 25-60 amino acids, such as of 26-60 amino acids, such as of 27-60 amino acids, such as of 28-60 amino acids, such as of 29-60 amino acids, such as of 30-60 amino acids, such as of 31-60 amino acids, such as of 32-60 amino acids, such as of 33-60 amino acids, such as of 34-60 amino acids, such as of 35-60 amino acids.
- 15 15. The isolated peptide according to any one of embodiments 1-14, which cell-penetrating peptide is of 18-60, such as 18-59, such as 18-58, such as 18-57, such as 18-56, such as 18-55, such as 18-54, such as 18-53, such as 18-52, such as 18-51, such as 18-50, such as 18-49, such as 18-48, such as 18-47, such as 18-46, such as 18-45, such as 18-44, such as 18-43, such as 18-42, such as 18-41, such as 18-40, such as 18-39, such as 18-38, such as 18-37, such as 18-35 amino acids, such as of 18-34 amino acids, such as of 18-33 amino acids, such as of 18-32 amino acids, such as of 18-31 amino acids, such as of 18-30 amino acids, such as of 18-29 amino acids, such as of 18-28 amino acids, such as of 18-27 amino acids, such as of 18-26 amino acids, such as of 18-25 amino acids, such as of 18-24 amino acids, such as of 18-23 amino acids, such as of 18-22 amino acids, such as of 18-21 amino acids, such as of 18-20 amino acids, such as of 18-19 amino acids.
- 20 25 16. The isolated peptide according to any one of embodiments 1-15, wherein the net charge of X^2 is below or equal to 0; and wherein X^1 and X^3 defines a sequence of 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof.
- 30

17. The isolated peptide according to any one of embodiments 1-16, wherein the net charge of X^2 is below or equal to 0; wherein the net charge of X^4 is above or equal to 1; and wherein X^1 and X^3 defines a sequence of 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof.
- 5 18. The isolated peptide according to any one of embodiments 1-17, wherein the net charge of X^2 and X^4 are below or equal to 0 and wherein X^1 and X^3 defines a sequence of 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof.
- 10 19. The isolated peptide according to any one of embodiments 1-18, wherein the net charge of X^2 and X^4 are above or equal to 1 and wherein X^1 and X^3 defines a sequence of 1 or 2 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof.
- 15 20. The isolated peptide according to any one of embodiments 1-19, wherein the net charge of X^2 is above or equal to 1; wherein the net charge of X^4 is below or equal to 0; wherein X^1 defines a sequence of 1 or 2 amino acids with a positive charge; and wherein X^3 defines a sequence of 2, 3 or 4 amino acid independently selected from any basic amino acid, citrulline, tryptophan, or a derivative thereof.
21. The isolated peptide according to any one of embodiments 1-20, wherein said basic amino acid is independently selected from Arg, Lys, and His.
- 20 22. The isolated peptide according to any one of embodiments 1-21, wherein X^2 and/or X^4 comprises the sequence of amino acids defined by position 135-157 of SEQ ID NO:3, or a fragment or variant thereof.
23. The isolated peptide according to any one of embodiments 1-22, wherein X^2 and/or X^4 consist of a sequence selected from GYIPLVGAPLG, GYLPAVGAPIG, GYLPAVGAPI, NYVTGNIPG, NYATGNLPG, NYATGNLPG, VTGNIPGSTYS, IRNLGRVIETLTG, SRNLGKVIDTLTC, IRNLGRVIETLT, GGGQIIGGNYLIP, GGGQIVGGVYLLP, LIFLARSALIV, LIFLARSALIL, LIFLARSALIL, SAYERMCNIL, SAYERZVNIL, TAYERZCNIL, IAYERMCNIL, IAYERMCNIL, LFFKCIYRLFHGL, LFFKTITRLFHGL, GLEPLVIAGILA, GSDPLVVAASIV, NLVPMVATV, NLVPMVATV, NIVPZVTA, PEVIPMFSALS, FIIPXFTALSG, ALGPAATL, GPVVHLTL, LECVYCKQQLL, GVDYAFRDLC, GVFDYAFRDIN, GATPVDLLGA, GVTPAGLIGV, VARALAHGVRV, VIRVIAHGLRL, GITFSIFLIVS, CSFSIFLLAL, GCSFSIFLLAL, GITFSIYLIVS, LZGYIPLIGA, LMGYIPLVGA, LZGYIPLIGA, PBIGVRATB, GPRLGVRATR, GPRLGVRAT, RGSVAHKS, SALILRGSAHKS, FQTAAQRAMM, FQTAAQRAVZ, FQTVVQBA, FQTAAQRA, GPSTEGVPESM,
- 25
- 30

LLSTEGVPNSZ, GSLVGLLHIVL, ASIVGILHLIL, NLVPMVATV, NIVPZVVT, TPQDLNMTLN,
ALLYGATPYAIG, MMTACQGVG, GQAGDDFS, EYDFAFRDLC, GFAFRDLCIVY, GFAYRDINLAY,
GALNLCLPM, GALQIBLPL, and IRNLGRVIETLTZGYIPLIGA, or a fragment or variant thereof.

24. The isolated peptide according to any one of embodiments 1-23, wherein X^2 and/or X^4
5 consist of a sequence derived from an amino acid sequence selected from SEQ ID NO:1, SEQ
ID NO:2, SEQ ID NO:3, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:6, SEQ ID NO:7, SEQ ID
NO:8, SEQ ID NO:9, SEQ ID NO:10, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, SEQ ID
NO:14, SEQ ID NO:15, SEQ ID NO:16, SEQ ID NO:17, SEQ ID NO:18, SEQ ID NO:19, SEQ
ID NO:20, SEQ ID NO:21, SEQ ID NO:22, SEQ ID NO:23, SEQ ID NO:24, SEQ ID NO:25,
10 SEQ ID NO:26, SEQ ID NO:27, SEQ ID NO:28, SEQ ID NO:29, SEQ ID NO:30, SEQ ID
NO:31, SEQ ID NO:32, SEQ ID NO:33, SEQ ID NO:34, SEQ ID NO:35, SEQ ID NO:36, SEQ
ID NO:37, SEQ ID NO:38, SEQ ID NO:39, SEQ ID NO:40, SEQ ID NO:41, SEQ ID NO:42,
SEQ ID NO:43, SEQ ID NO:44, SEQ ID NO:45, SEQ ID NO:46, SEQ ID NO:126, and SEQ ID
NO:198, or a fragment or variant thereof.
- 15 25. The isolated peptide according to any one of embodiments 1-24, wherein said peptide
is selected from RRGYIPLVGAPLGBGRVARALAHGVRV (SEQ ID NO:47),
RGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:48), RGYIPLVGAPLGRRRVARALAHGVRV (SEQ
ID NO:49), RRGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:50),
RRGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:51), BRGYIPLVGAPLGRRRVARALAHGVRV
20 (SEQ ID NO:52), RRRGYIPLVGAPLGBRVARALAHGVRV (SEQ ID NO:53),
RGYIPLVGAPLGKKKVARALAHGVRV (SEQ ID NO:54), RGYIPLVGAPLGRRRVARALAHGVRV (SEQ
ID NO:55), KKGYIPLVGAPLGKKKVARALAHGVRV (SEQ ID NO:56),
WGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:57), WWGYIPLVGAPLGRRRVARALAHGVRV (SEQ
ID NO:58), EEGYIPLVGAPLGEEVARALAHGVRV (SEQ ID NO:59),
25 GGGYIPLVGAPLGGGVARALAHGVRV (SEQ ID NO:60), EEGYIPLVGAPLGRRRVARALAHGVRV (SEQ
ID NO:61), RRGYIPLVGAPLGLRRVARALAHGVRV (SEQ ID NO:62),
WWGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:63), WWGYIPLVGAPLGRRRVARALAHGVRV
(SEQ ID NO:64), WWGYIPLVGAPLGRVARALAHGVRV (SEQ ID NO:65),
RGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:66), RRGYLPVAVGAPIGBRVIRVIAHGLRL (SEQ ID
30 NO:67), RRGYIPLVGAPLGBRVARALAHGVRV (SEQ ID NO:68), GYIPLVGAPLGGVARALAHGVRV
(SEQ ID NO:69), WWGYLPVAVGAPIRRVIRVIAHGLRL (SEQ ID NO:70),
GYIPLVGAPLGGVARALAHGVRV (SEQ ID NO:71), RRGYIPLVGAPLGBGRVARALAHGVRV (SEQ ID
NO:72), RGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:73),
RGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:74), RRGYIPLVGAPLGRRRVARALAHGVRV (SEQ
35 ID NO:75), RRGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:76),
BRGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:77), RRRGYIPLVGAPLGBRVARALAHGVRV
(SEQ ID NO:78), RGYIPLVGAPLGKKKVARALAHGVRV (SEQ ID NO:79),

RGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:80), KKGYIPLVGAPLGKKVARALAHGVRV (SEQ ID NO:81), WGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:82),
 WWGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:83), RRGYIPLVGAPLGRRRVARALAHGVRV (SEQ ID NO:84), RRNYVTGNIPGBRGITFSIFLIVS (SEQ ID NO:85),
 5 WWNYATGNLPGRRCSFSIFLLAL (SEQ ID NO:86), WWNYVTGNIPGBRGITFSIFLIVS (SEQ ID NO:87), WWNYVTGNIPGRRGITFSIFLIVS (SEQ ID NO:88), RRNYATGNLPGRRGCSFSIFLLAL (SEQ ID NO:89), RRVTGNIPGSTYSGBRGITFSIYLIVS (SEQ ID NO:90),
 RRIRNLGRVIETLTGBRLZGYIPLIGA (SEQ ID NO:91), RRSRNLGKVIDTLTCBRLMGYIPLVGA (SEQ ID NO:92), SRNLGKVIDTLTCGFADLMGYIPLVGA (SEQ ID NO:93),
 10 WWIRNLGRVIETLTRRLZGYIPLIGA (SEQ ID NO:94), WWSRNLGKVIDTLTCRRLMGYIPLVGA (SEQ ID NO:95), RRGGGQIIGGNYLIPRBPBIVRATB (SEQ ID NO:96),
 GGGQIVGGVYLLPRRGPRLGVRATR (SEQ ID NO:97), RRGGGQIVGGVYLLPRRGPRLGVRATR (SEQ ID NO:98), WWGGGQIVGGVYLLPRRGPRLGVRAT (SEQ ID NO:99), BRILFLARSALIVRGSLVAHKS (SEQ ID NO:100), EDLIFLARSALILRGSLVAHKS (SEQ ID NO:101),
 15 BRILFLARSALILBGRSALILRGSLVAHK (SEQ ID NO:102), SAYERMCNILKGKFQTAAQRAMM (SEQ ID NO:103), SAYERZVNILKGKFQTAAQRAVZ (SEQ ID NO:104),
 BRTAYERZCNILBRGRFQTVVQBA (SEQ ID NO:105), BRIAYERMCNILLBRGKFQTAAQRA (SEQ ID NO:106), IAYERMCNILKGKFQTAAQRA (SEQ ID NO:107), LFFKCIYRLFKHGLKRGPPSTEGVPESM (SEQ ID NO:108), BRRLFFKTITRLFBHGLRRLSTEGVPNSZ (SEQ ID NO:109),
 20 BRGLEPLVIAGILARRGSLVGLLHIVL (SEQ ID NO:110), BRGSDPLVVAASIVRRASIVGILHLIL (SEQ ID NO:111), RNLVPMVATVRRNLVPMVATVB (SEQ ID NO:112), RNLVPMVATVBRRNLVPMVATVB (SEQ ID NO:113), RNIVPZVVTARRNIVPZVVTAB (SEQ ID NO:114),
 PEVIPMFSALESGATPQDLNMTLN (SEQ ID NO:115), RFIIPXFTALSGRRALLYGATPYAIG (SEQ ID NO:116), KALGPAATLEEMMTACQGVG (SEQ ID NO:117), RRGPPVHLLTLLRRRGQAGDDFS (SEQ ID NO:118), RRGPPVHLLTLLRRRGQAGDDFS (SEQ ID NO:119), RRGPPVHLLTLLRRRGQAGDDFS (SEQ ID NO:120), RRLECVYCKQQLLRREVYDFAFRDLC (SEQ ID NO:121),
 25 RRGVYDFAFRDLCRRGFAFRDLCIVYR (SEQ ID NO:122), RRGVFDYAFRDINRRGFAYRDINLAYR (SEQ ID NO:123), RRGATPVDLLGARRGALNLCPLMR (SEQ ID NO:124),
 RRGVTPAGLIGVRRGALQIBLPLR (SEQ ID NO:125), RGYLPAVGAPIGRRRVIRVIAHGLRLR (SEQ ID NO:196), RRSRNLGKVIDTLTCRRLMGYIPLVGA (SEQ ID NO:197), and
 30 RRIRNLGRVIETLTZGYIPLIGARRIRNLGRVIETLTZGYIPLIGAR (SEQ ID NO:199), or a fragment or variant thereof.

26. The isolated peptide according to any one of embodiments 1-25, wherein the peptide consist of a sequence selected from X¹-NYVTGNIPG-X³-GITFSIYLIVS; X¹-IRNLGRVIETLT-X³-LZGYIPLIGA; X¹-GYLPAVGAPI-X³-VIRVIAHGLRL; X¹-GGGQIIGGNYLIP-X³-PBIVRATB; X¹-NYATGNLPG-X³-GCSFSIFLLAL; X¹-SRNLGKVIDTLTC-X³-LMGYIPLVGA; X¹-GYIPLVGAPL-X³-VARALAHGVRV; X¹-GGGQIVGGVYLLP-X³-PRLGVRATR; X¹-LTFLVRSVLLI-X³-GSVLIVRGSLVH; X¹-TAYERZCNIL-X³-GRFQTVVQBA; X¹-SDPLVVAASIV-X³-ASIVGILHLIL; X¹-LIFLARSALIL-X³-

- SALILRGSAH; X¹-IAYERMENIL-X³-GKFQTAAQRA; and X¹-LEPLVIAGILA-X³-GSLVGLLHIVL; X¹-NLVPMVATV-X³-NLVPMATV; X¹-GYLPAVGAPIG-X³-VIRVIAHGLRL; X¹-IRNLGRVIETLTG-X³-LZGYIPLIGA; X¹-GVYDFAFRDLC-X³-GFAFRDLCIVYR, X¹-GVFDYAFRDIN-X³-GFAYRDINLAYR, X¹-GATPVDLLGA-X³-GALNLCPLMR, X¹-GVTPAGLIGV-X³-GALQIBLPLR, and X¹-
- 5 IRNLGRVIETLTZGYIPLIGA-X³-IRNLGRVIETLTZGYIPLIGA; optionally with an X⁵ in the C-terminal of the peptide, wherein X¹ and X³ and X⁵ refers to X¹, X³, and X⁵ of formula I.
27. The isolated peptide according to any one of embodiments 1-26, wherein the peptide comprises one or more cysteine.
28. The isolated peptide according to any one of embodiments 1-27, wherein the peptide
10 contain intramolecular bonds, such as intramolecular disulfide (S-S) bonds between two cysteine residues or acylals.
29. The isolated peptide according to any one of embodiments 1-28, wherein the N- and/or C-terminal amino acid in X² is a hydrophilic or polar amino acid.
30. The isolated peptide according to any one of embodiments 1-29, wherein the N-
15 terminal amino acid in X⁴ is a hydrophilic or polar amino acid.
31. The isolated peptide according to any one of embodiments 1-30, which is not more than 58 amino acids, such as not more than 56, 54, 52, 50, 48, 46, 44, 42, 40, 38, 36, 34, 32, 30, 28, 26, 24, 22, 20, 18 amino acid residues.
32. The isolated peptide according to any one of embodiments 1-31, wherein X¹ consist of
20 2 or 3 amino acids.
33. The isolated peptide according to embodiment 32, wherein X¹ consist of WW, BR, or RR.
34. The isolated peptide according to any one of embodiments 1-33, wherein X³ consist of 2 or 3 amino acids.
- 25 35. The isolated peptide according to embodiment 34, wherein X³ consist of WW, BR, or RR.
36. Isolated peptide of not more than 60 amino acids comprising a sequence of X² and/or X⁴ as independently defined in any one of table 1 or table 2.

37. Isolated peptide according to embodiment 36, which peptide consists of a sequence of X^2 or X^4 as independently defined in any one of table 1 or table 2.
38. The isolated peptide according to any one of embodiments 1-37, wherein X^2 defines a sequence of 8-25 amino acids, such as 8-20 amino acids, such as 8-15 amino acids.
- 5 39. The isolated peptide according to any one of embodiments 1-38, wherein X^4 defines a sequence of 8-25 amino acids, such as 8-20 amino acids, such as 8-15 amino acids.
40. The isolated peptide according to any one of embodiments 1-39, wherein X^2 defines a sequence of less than 25, such as less than 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7 or 6 amino acids.
- 10 41. The isolated peptide according to any one of embodiments 1-40, wherein X^2 defines a sequence of more than 8, such as more than 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29 amino acids.
42. The isolated peptide according to any one of embodiments 1-41, wherein X^4 defines a sequence of less than 25, such as less than 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13,
15 12, 11, 10, 9, 8, 7 or 6 amino acids.
43. The isolated peptide according to any one of embodiments 1-42, wherein X^4 defines a sequence of more than 8, such as more than 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29 amino acids.
44. The isolated peptide according to any one of embodiments 1-43, wherein the cell-
20 penetrating peptide of the invention does not consist of the following sequence
RFIIP[Nle]FTALSGGRRALLYGATPYAIG, where Nle denotes a nor-leucine.
45. The isolated peptide according to any one of embodiments 1-44, wherein X^2 and/or X^4 is not derived from HIV.
46. The isolated peptide according to any one of embodiments 1-45, wherein X^4 is a linear
25 sequence of less than 12 amino acids.
47. The isolated peptide according to any one of embodiments 1-46, wherein X^2 is a linear sequence of less than 12 amino acids.

48. The isolated peptide according to any one of embodiments 1-47, wherein X^2 and/or X^4 do not contain nor-leucine.
49. The isolated peptide according to any one of embodiments 1-48, wherein X^2 do not contain nor-leucine.
- 5 50. The isolated peptide according to any one of embodiments 1-49, wherein X^2 and/or X^4 only contains natural amino acids.
51. The isolated peptide according to any one of embodiments 1-50, wherein X^2 only contains natural amino acids.
52. The isolated peptide according to any one of embodiments 1-51, wherein X^2 only
10 contains natural amino acids if derived from HIV.
53. The isolated peptide according to any one of embodiments 1-52, wherein X^2 and/or X^4 is derived from HCV, CMV, HPV, Influenza, adenoviruses, or picornaviruses.
54. Use of a peptide comprising a sequence of X^2 or X^4 as independently defined in any one of table 1 or table 2 for inducing an immune response in a subject.
- 15 55. A dimer peptide comprising two peptides monomers, wherein each peptide monomer is as defined in any one of embodiments 1-53.
56. A dimer peptide according to embodiment 55, wherein the two peptide monomers are identical.
57. Peptide combination comprising two or more peptide according to any one of
20 embodiments 1-53.
58. An isolated nucleic acid or polynucleotide encoding a peptide according to any one of embodiments 1-53.
59. A vector comprising the nucleic acid or polynucleotide according to embodiment 58.
60. A host cell comprising the vector according to embodiment 59.

61. An immunogenic composition comprising at least one peptide according to any one of embodiments 1-53, a dimer peptide according to any one of embodiments 55-56, a peptide combination according to embodiment 57, the nucleic acid or polynucleotide according to embodiment 58, or the vector according to embodiment 59; in combination with a
5 pharmaceutically acceptable diluent or vehicle and optionally an immunological adjuvant.

62. The immunogenic composition according to embodiment 61 in the form of a vaccine composition.

63. A method for inducing an immune response in a subject against an antigen which comprises administration of at least one peptide according to any one of embodiments 1-53, a dimer peptide according to any one of embodiments 55-56, a peptide combination
10 according to embodiment 57, the nucleic acid or polynucleotide according to embodiment 58, or the vector according to embodiment 59; or the composition according to any one of embodiments 61-62.

64. A method for reducing and/or delaying the pathological effects of a virus in a subject infected with said virus, the method comprising administering an effective amount of at least one peptide according to any one of embodiments 1-53, a dimer peptide according to any one of embodiments 55-56, a peptide combination according to embodiment 57, the nucleic acid or polynucleotide according to embodiment 58, or the vector according to embodiment 59; or the composition according to any one of embodiments 61-62.
15

65. A peptide according to any one of embodiments 1-53 for use as a medicament.
20

66. A peptide according to any one of embodiments 1-53 for treating the pathological effects of a virus in a subject infected with said virus.

Sequence list (amino acids in bold represents suitable antigenic sequences that may be used as any of X² and/or X⁴ as defined in formula I of the present invention)

25 SEQ ID NO:1: Accession no AF009606; Hepatitis C virus subtype 1a polyprotein gene, complete cds.

30 MSTNPKPQRKTKRNTNRRPQDVKFPGGGQIVGGVYLLPRRGPRL
GVRATRKTSESRQPRGRRQPIPKARRPEGRTWAQPGYPWPPLYGNEGCGWAGWLLSPRG
SRPSWGPTDPRRRSRNLGKVIDTLTCGFADLMGYI**PLVGAPLGGAARALAHGVRVLED**
GVNYATGNLPGCSFSIFLLALLSCLTVPASAYQVRNSSGLYHVTNDCPNSSIVYEAAD
AILHTPGCVPCVREGNASRCWVAVTPTVATRDKLPTTQLRRHIDLLVGSATLCSALY
35 VGDLCGSVFLVGQLFTFSRRHWTQDCNCSTYPGHITGHRMAWMMMNWSPTAALVV
AQLLRIPQAIMDMIAGAHWGVLAGIAYFSMVGNWAKVLVLLLFAGVDAETHVTGGSA
GRRTAGLVGLLTPGAKQNIQLINTNGSWHINSTALNCNESLNTGWLAGLFYQHKFNSS
GCPERLASCRRLTDFAGWGPISYANGSGLDERPYCWHYPPRPGCIVPAKSVCGPVC

- 5 FTSPVVGTTDRSGAPTYSWGANDTDVFLNNTTRPPLGNWFGCTWMNSTGFTKVCGA
 PPCVIGGVGNNTLLCPTDCFRKHPEATYSRCGSGPWITPRCMVDYPYRLWHYPCTINY
 TIFKVRMYVGGVEHRLEAACNWTRGERCDLEDRDRSELSPLLLSTTQWQVLPSCSFTTL
 PALSTGLIHLHQINVDVQYLYGVGSSIASWAIKWEYVVLFLLLADARVCSCLWMMLL
 10 ISQAEAALENLVI LNAASLAGTHGLVSFLVFFCFAWYLKGRWVPGAVYAFYGMWPLLL
 LLLALPQRAYALDTEVAASCGGVVLVGLMALTLSPYYKRYISWCMWWLQYFLTRVEAQ
 LHVWVPPPLNVRGGRDAVILLMCVVHPTLVFDITKLLLAIFGPLWILQASLLKVPYFVR
 VQGLLRICALARKIAGGHYVQMAIIKLGALTGTYYVNHLPDRDWAHNGLRDLAVAVE
 PVVFSRMETKLITWGADTAACGDIINGLPVSARRGQEIILGPADGMVSKGWRLAPIT
 15 AYAQQTRGLLGCIITSLTGRDKNQVEGEVQIVSTATQTFLATCINGVCWTVYHGAGTR
 TIASPKGPVIMYTNVDQDLVGWPAPQGSRLTPCTCGSSDLYLVTRHADVIPVRRRG
 DSRGSLSPRISYLGSSGGPLLCPAGHAVGLFRAAVCTRGVAKAVDFIPVENLETT
 MRSFVFTDNSSPPAVPQSFQVAHLHAPTGSKGSTKVPAAYAAQGYKVLVLPNSVAATL
 GFGAYMSKAHGVDPNIRTGVRTITTTGSPITYSTYKGFLADGGCSGGAYDIIICDECHS
 20 TDATSI LGIGTVLDQAETAGARLVV LATATPPGSVTVSHPNIEEVALSTTGEIPFYGK
 AIPLEVIKGRHLLFCHSKKKCELA AKLVALGINAVAYYRGLDVSVIPTSGDVVVVS
 TDALMTGFTGDFDSDVDCNTCVTQTVDFSLDPTFTIETTLTPQDAVSRTQRRGRTGRG
 KPGIYRFVAPGERP SGMFDSVLCECYDAGCAWYELT PAETTVRLRAYMNT PGLPVCQ
 DHLEFWEGVFTGLTHIDAHFLSQTKQSGENFPYLVAYQATVCARAQAPPSWDQMWC
 25 LIRLKPTLHGPTPLLYRLGAVQNEVT LTHPITKYIMTCMSADLEVVTSTWVLVGGVLA
 ALAAYCLSTGCVVIVGRIVLSGKPAIIPDREVLYQEFDEMEEC SQHLPIYIEQGMMLAE
 QFKQKALGLLQTSARQA EVITPAVQTNWQKLEVFWAKHMMNFI SGIQYLAGLSTLPGN
 PAIASLMAFTA AVTSPLTTGQTLFNILGGWVAAQLAAPGAATAFVGAGLAGAAIGSV
 GLGKVLVDILAGYGAGVAGALVAFKIMSGEVPSTEDLVNLLPAILSPGALVVGVCVAA
 30 I LRRHVGPGEAGVQWMNRLIAFASRGNHVSPHYVPESDAAARVTAI LSSLTVTQLLR
 RLHQWISSECTTPCSGSWLRDIWDWICEVLSDFKTWLKAKLMPQLPGIPFVSCQRGYR
 GVWRGDGIMHTRCHCGAEITGHVKNGTMRIVGPRTCRNMWSGTFFINAYTTGPCTPLP
 APNYKFALWRVSAEEYVEIRRVGDFHYVSGMTTDNLKCPQIPSPFEFFTELDGVRLLHR
 FAPPCKPLLEEVSVFRVGLHEYFVGSQLPCEPEPDVAVLTSMLTDP SHITAEAAGRRL
 35 ARGSPPSMASSAQSAPSLKATCTANHDS PDAELIEANLLWRQEMGGINTRVESEN
 KVVILDSFDPLVAEEDEREVSVP AEILRKSRRFARALPVWARP DYNPPLVETWKKPDY
 EPPVHGCPLPPRSPVPVPPRKKRTVVLTESTLSTALAE LATSFGSSSTSGITGDN
 TTTSSEAPSGCPPDS DVESYSSMPPLEGE PGDPDLSDGSWSTVSSGADTEDVCCSM
 SYSWTGALVTPCAAEEQKL PINALSNSLLRHHNLVYSTTSRSACQRQKKVTFDRLQVL
 40 DSHYQDVLKEVKAAASKVKANLLSVEEACSLTPPHSAKS KFGYGAKDVRCHARKAVAH
 INSVWKDLLED SVTPIDTTIMAKNEVFCVQPEKGGKRKPARLIVFPDLGVRVCEKMALY
 DVVSKLPLAVMGSSYGFQYSPGQ RVEFLVQAWKS KPTPMGFSYDTRCFDSTVTESDIR
 TEEAIYQCCDLDPQARVAIKSLTERLYVGGPLTNSRGENC GYRRCRASGVLT TSCGNT
 LTCYIKARAACRAAGLQDCTMLVCGDDL VICESAGVQEDAASLRAFTEAMTRY SAPP
 45 GPPQPEYDLELITSCSSNVSAHDGAGKRVY YLTRDPTPLARA AWETARHTPVNSW
 LGNIIMFAPTLWARMILMTHFFSVLIARDQLEQALNCEIYGACYSIEPLDLPPIIQR L
 HGLSAFSLHSYSPGEINRVAACLRKLGVPPLRAWRHRARSVRARLLSRGGRAAICGKY
 LFNWAVRTKLKLTPIAAAGRLDLSGWFTAGYSGGDIYHSVSHARPRWF FCLLLAAG
 VGIYLLPNR
- 50 SEQ ID NO:2: HCV core protein, H77, Accession AF009606
 Genbank number: 2316097
 >gi|2316098|gb|AAB66324.1| polyprotein [Hepatitis C virus subtype 1a]
 MSTNPKPQRKTKRNTNRRPQDVKFPGGGQIVGGVYLLPRRGPRLGVRATRKTSERSQPRGRRQPIPKARR
 PEGRTWAQPGYPWPPLYGNEGCWAGWLLSPRGSRPSWGPDPFRRRSRNLGKVIDTLTCGFADLMGYIPLVGAPLGGAAR
 ALAHGVRVLEDGVNYATGNLPGCSFSIFLLALLSCLTVPASA
- 55 SEQ ID NO:3:
 Hepatitis C virus mRNA, complete cds; ACCESSION M96362 M72423; Hepatitis C
 virus subtype 1b
 MSTNPKPQRKTKRNTNRRPQDIKFPGGGQIVGGVYLLPRRGPRL
 GVRATRKTSERSQPRGRRQPIPKARRPEGRAWAQPGYPWPPLYGNEGLGWAGWLLSPRG
 SRPSWGPDPFRRR**SRNLGKVIDTLTCGFADLMGYIPLVGAPLGVARALAHGVRVLED**
 60 **GVNYATGNLPGCSFSIFLLALLSCLTTPVSAYEVRNASGMYHVTNDCSNSSIVYEAD**
 MIMHTPGCVPCVREDNSSRCWALTPTLAARNASVPTTTLRRHVDLLVGVAAFCSAMY
 VGDLCGSVFLVSQLFTFSPPRRHETVQDCNCSIYPGRVSGHRMAWDMMNWSPTTALVV

5 SQLLRIPQAVVDMVTGSHWGI LAGLAYYSMVGNWAKVLIAMLLFAGVDGTHVTGGAQ
 GRAASSLTSLFSPGPVQHLQLINTNGSWHINRTALSCNDSLNTGFVAALFYKYRFNAS
 GCPERLATCRPIDTFAQGWGPITYTEPHDLQDRPYCWHYAPQPCGIVPTLQVCGPVYC
 10 FTPSPVAVGTTDRFGAPTYRWGANETDVLLLNAGPPQGNWFGCTWMNGTGFTKTCGG
 PPCNIGGVGNNTLTCTDCFRKHPGATYTKCGSGPWLTPRCLVDYPYRLWHYPCTVNF
 TIFKVRMYVGGAEHRLDAACNWRGERCDLEDRDRSELSPLLLSTTEWQVLPSCFTTL
 PALSTGLIHLHQNIVDIQYLYGIGSAVVSFAIKWEYIVLLFLLLADARVCACLWMLL
 VAQAEAALENLVVNAASVAGAHGILSFIVFFCAAWYIKGRVPGAAYALYGVWPLLL
 15 LLLALPPRAYAMDREMAASCGGAVFVGLVLLTSLPHYKVFLARFIWWLQYLITRTEAH
 LQVWVPLNVRGGRDAIILLTCVVHPELIFDITKYLLAIFGPLMVLQAGITRVFYFVR
 AQGLIRACMLARKVVGHHYQVMFMKLAALAGTYVVDHLTPLRDWAHTGLRDLAVAVE
 PVVFSDMETKVI TWGADTAACGDIILALPASARRGKEILLGPADSLEGQGWRLAPIT
 AYSQQTGRLLGCIITSLTGRDKNQVEGEVQVSTATQSFLATCINGVCWTVFHHGAGSK
 20 TLAGPKGPITQMTYNVDQDLVGPAPPGARSLTPCTCGSSDLYLVTRHADVIPVRRRG
 DGRGSLPPRPVSYLKGS SGGPLLCPSGHAVGILPAAVCTRGVAMAVEFIPVESMETT
 MRSPVFTDNPSPPAVPQTQVAHLHAPTGS GKSTRVPAAYAAQGYKVLVLPNSVAATL
 FGAYMSKAHGIDPNLRTGVRTITTGAPITYSTYGFADGGSGGAYDIIMCDECHS
 TDSTTIYIGITGVLDAQETAGARLVVLSTATPPGSVTVPHLNIIEVALSNTGEIPFYGK
 25 AIPTEAIKGRHILFCHSKKKCELAALKSLGLGLNAVAYYRGLDVSVIPTSGDVVVVA
 TDALMTGFTGDFDSVIDCNTCVTQTVDFSLDPTFTIETTVPQDAVSRQRGRGRTGRG
 RAGIYRFVTPGERPSGMFDSVLCECYDAGCAWYELTPAETSVRLRAYLNTPLGPVCQ
 DHLEFSEGVFTGLTHIDAHFSLQTKQAGENFPYLVAYQATVCARAQAPPSWDEMWR
 LIRLKPTLHGPTPLLYRLGAVQNEVTLTHPITKFIMTCMSADLEVVTSTWVLVGGVLA
 30 ALAAAYCLTTGSGVIVGRIILSGKPAIIPDREVLYQEFDEMEECASHLPYFEQGMQLAE
 QFKQKALGLLQATKQAEAAAPVVESKWRALETFWAKHMWNFISGIQYLAGLSTLPGN
 PAIRSPMAFTASITSPLTQTHTLLFNILGGWVAAQLAPPSAASAFVGAGIAGAAGVTI
 GLGKVLVDILAGYGAGVAGALVAFKIMSGEMPSAEDMVNLLPAILSPGALVVGIVCAA
 ILLRHVGPGEAGVQWMNRLIAFASRGNHVS PRHYVPESEPAARVTQIILSSLTITQLLK
 35 RLHQWINEDCSTPCSSSWLREIWDWICTVLTDFKTWLQSKLLPRLPGVPPFFSCQRGYK
 GVWRGDGIMHTTCPCGAQITGHVKNKSMRIVGPKTCSNTWYGTFFPINAYTTGPCTPSP
 APNYSKALWRVAAEEYVEVTRVGDFHYVTGMTTDNVKCPQVPAPEFFTEVDGVRLLHR
 YAPACRPLLLREEVVFQVGLHQYLVGSQLPCEPEPDVAVLTSMLTDP SHITAETA KRRL
 ARGSPPSLASSASQLSAPSLKATCTTHHDSPADLIEANLLWRQEMGGNITRVESEN
 40 KVVILDSFDPLRAEDDEGEISVPAEILRKS RKFPPALPIWAPPDYNPPLLESWKDPDY
 VPPVHGCPLPPTKAPPIPPPRRKRTVVLTESTVSSALAE LATKTFGSSGSSAIDSGT
 ATAPPDQASGDGDRES DVESFSSMPPLEGE PGDPDLS DGSWSTVSEEASEDVVCCSMS
 YTWGTALITPCAAEESKLPINPLSNSLLRHHNMVYATTSRSAGLRQKKVTFDRLQVLD
 DHYRDVLKEMKAKASTVKAKLLSVEEACKLTPPHSAKSKFGYGAKDVRSLSRAVTHI
 45 RSVWKDILLEDTE TETPISTTIMAKNEVFCVQPEKGRKPARLIVFPDLGVRVCEKMALYD
 VVSTLPQAVMGSSYGFQYSPKQ RVEFLVNTWKS KKCMPMGFSYDTRCFDSTVTENDIRV
 EESIYQCCDLAPEAKLAIKSLTERLYIGGPLTNSKGQNCGYRRCRASGVLTTSCGNTL
 TCYLKATAACRAAKLRDCTMLVNGDDLVI CESAGTQEDAASLRVFTTEAMTRY SAPPG
 DPPQPEYDLELITSCSSNVSVAH DASGKRVYLT RDPTT PLARA AWETARHTPVNSWL
 50 GNIIMYAPTLWARMILMTHFFSILLAQEQLEKTLDCQIYGACYSIEPLDLPQIIERLH
 GLSAFSLHSYSPGEINRVASCLRKLGVPPLRAWHRARSVRAKLLSQGGRAATCGKYL
 FNWAVRTKLKLTPIPAASRLDLSGW FVAGYSGGDIYHSLSRARPRWFMLC LLLLSVGV
 GIYLLPNR

50

SEQ ID NO:4, nucleocapsid protein of influenza A virus

1 MASQGTKRSY EQMETS GERQ NATEIRASVG RMVGGIGRFY IQMCTELKLS DHEGRLIQNS
 61 ITIERMVL SA FDERRNKYLE EHPSAGKDPK KTG GP IYRRR DGKWMRELIL YDKEEIRRIW
 121 RQANNGEDAT AGLTHMMIWH SNLNDATYQR TRALVRTGMD PRMC SLMQGS TLPRRSGAAG
 55 181 AAVKGVGTMV MELIRMIKRG INDRNFWRGE NGRRTRIAYE RMCN ILKGKF QTAAQRAMMD
 241 QVRESRNP GN AEIEDLFLA RSALILRGSV AHKSCLPACV YGLAVASGYD FEREGYSLVG
 301 IDPFRLQNS QVFSLRPNE NPAHKSQLVW MACHSAAFED LRVSSFIRGT RVVPRGQLST
 361 RGVQIASNEN METMDSSTLE LRSRYWAIRT RSGGNTNQQR ASAGQISVQP TFSVQRNLFP
 421 ERATIMAAFT GNTEGRTS DM RTEIIRMMEN ARPEDVSFQG RGVFELSDEK ATNP IVP SFD
 60 481 MSNEGS

SEQ ID NO:5

>gi|73919153|ref|YP_308840.1| matrix protein 2 [Influenza A virus (A/New York/392/2004 (H3N2))]

5 MSLLTEVETPIRNEWGCRCNDS**SDPLVVAASIIGILHLILWILDR**LFFKCVYRLFKHGLKRG**SPSTEGVPE** 70
SMREEYRKEQQNAVDADDSHFVSIELE

SEQ ID NO:6

10 >gi|73919147|ref|YP_308843.1| nucleocapsid protein [Influenza A virus (A/New York/392/2004 (H3N2))]
MASQGTKRSYEQMETDGDQRNATEIRASVGKMDIGIGRFYIQMCTELKLS**DHEGR**LIQNSLTIEKMVLSA 70
FDERRNKYLEEHP**SAGKDPKKTGGPIYRRVDGKWMREL**VLYDKEEIRRIWRQANNGEDATAGLTHIMIWH 140
SNLN**DATYQ**TRALV**RTGMDPRMCSLMQGSTLP**RRSGAAGAAVKGIGTMV**MELIR**MV**KRGINDR**NFWRGE 210
NGRKTR**SAYERMCN**ILKGK**FQTAAQ**RAMVDQVRESRNP**GNAEIEDLI**FLARSALIL**RGSVAHKS**CLPACA 280
15 YGPAVSSGYD**FEKEGYSLVGIDPFKLLQNSQIYSLIRPNEN**PAHKSQ**LVWMACHS**AAFEDLRLLSFIRGT 350
KVSPRGKLSTRGVQ**IASNENMDNMGSSTLELR**SGYWAI**TRSGGNTNQQRASAGQTSVQPTFSVQRNL**PF 420
EKSTIMAAFTGNTEGRTSDMR**AEIIRMEGAKPEEVSFRGRGVFELSDEKATNP**IVPSFDMSNEGSYFFG 490
DNAEEYDN
--

20 SEQ ID NO:7
>gi|56583270|ref|NP_040979.2| matrix protein 2 [Influenza A virus (A/Puerto Rico/8/34 (H1N1))]
25 MSLLTEVETPIRNEWGCRCNGSS**SDPLAIAANIIGILHLILWILDR**LFFKCIYRRFKYGLK**GGPSTEGVPK**
SMREEYRKEQQSAVDADDGHFVSIELE

SEQ ID NO:8
>gi|8486130|ref|NP_040982.1| nucleocapsid protein [Influenza A virus (A/Puerto Rico/8/34 (H1N1))]
30 MASQGTKRSYEQMETDGERQ**NATEIRASVGK**MDIGIGRFYIQMCTELKLS**DYEGR**LIQNSLTIERMVLSA
FDERRNKYLEEHP**SAGKDPKKTGGPIYRRVNGKWMREL**LILYDKEEIRRIWRQANNGDDATAGLTHMMIWH
SNLN**DATYQ**TRALV**RTGMDPRMCSLMQGSTLP**RRSGAAGAAVKGVGT**VMELVRMI**KRGINDRNFWRGE
NGRKTR**IAYERMCN**ILKGK**FQTAAQ**KAMMDQVRESRDP**GNAEFEDLT**FLARSALIL**RGSVAHKS**CLPACV
35 YGPAVASGYD**FEREGYSLVGIDPFRL**LQNSQVYSLIRPNENPAHKSQ**LVWMACHS**AAFEDLRVLSFIKGT
KVSPRGKLSTRGVQ**IASNENMETMESSTLELR**SYWAI**TRSGGNTNQQRASAGQISIQPTFSVQRNL**PF
DRTT**VMAAFTGNTEGRTSDMRTEIIRME**SAR**PEDVSFQGRGVFELSDEKAASPIVPSFDMSNEGSYFFG**
DNAEEYDN
--

40 SEQ ID NO:9
>gi|73912687|ref|YP_308853.1| membrane protein M2 [Influenza A virus (A/Korea/426/68 (H2N2))]
MSLLTEVETPIRNEWGCRCNDS**SDPLVVAASIIGILHFI**LWILDR**LFFKCIYRFFKHGLKRG**SPSTEGVPE
45 **SMREEYRKEQQSAVDADD**SHFVSIELE

SEQ ID NO:10
>gi|73921307|ref|YP_308871.1| nucleoprotein [Influenza A virus (A/Korea/426/68 (H2N2))]
50 MASQGTKRSYEQMETDGERQ**NATEIRASVGK**MDIGIGRFYIQMCTELKLS**DYEGR**LIQNSLTIERMVLSA
FDERRNKYLEEHP**SAGKDPKKTGGPIYKRVDGKWMREL**VLYDKEEIRRIWRQANNGDDATAGLTHMMIWH
SNLN**DTTYQ**TRALV**RTGMDPRMCSLMQGSTLP**RRSGAAGAAVKGVGT**VMELIRMI**KRGINDRNFWRGE
NGRKTR**SAYERMCN**ILKGK**FQTAAQ**RAMMDQVRESRNP**GNAEIEDLI**FLARSALIL**RGSVAHKS**CLPACV
YGPAIASGYN**FEKEGYSLVGIDPFKLLQNSQVYSLIRPNEN**PAHKSQ**LVWMA**CNSAAFEDLRVLSFIKGT
55 KVSPRGKLSTRGVQ**IASNENMDTMESSTLELR**SYWAI**TRSGGNTNQQRASAGQISVQPAF**SVQRNL**PF**
DKPTIMAAFTGNTEGRTSDMR**AEIIRMEGAKPEEMS**FQGRGVFELSDEKATNP**IVPSFDMSNEGSYFFG**
DNAEEYDN

SEQ ID NO:11
 >gi|330647|gb|AAA45994.1| pp65 [Human herpesvirus 5]
 MASVLGPISGHVLKAVFSRGDTPVLPHETRLLQTGIHVRVSQPSLILVSQYTPDSTPCHRGDNQLQVQHT 70
 YFTGSEVENVSVNVHNPTGRSICPSQEPMSIYVYALPLKMLNIP SINVHHYPSAAERKHRHLPVADAVIH 140
 5 ASGKQMWQARLTVSGLAWTRQQNQWKEPDVYYTSAFVFPTKDVALRHVVCAHELVCSEMENTRATKMQVIG 210
 DQYVKVYLESFCEDEVPSGKLFMHVTLGSDVEEDLTMTRNQPFMRPHERNGFTVLCPKNMIKPGKISHI 280
 MLDVAFTSHEHFGLLCPKSIPLGLSISGNLLMNGQQIFLEVQAIRETVELRQYDPVAALFFFDIDLLLQRG 350
 PQYSEHPTFTSQYRIQGKLEYRHTWDRHDEGAAGDDVWTSGSDSDEELVTTERKTTPRTGGGAMAGAS 420
 TSAGRKRKSASSATACTAGVMTRGRLKAESTVAPEEDTDESDSNEIHNPVFTWPPWQAGILARNLVPMV 490
 10 **ATV**QGQNLKYQEFFFWDANDIYRIFAELEGVWQPAAQPKRRRHRQDALPGPCIASTPKKHRG 541

SEQ ID NO:12
 >gi|33330937|gb|AAQ10712.1| putative transforming protein E6 [Human
 15 papillomavirus type 16]
 MHQKRTAMFQDPQERPGKLPQLCTELQTTIHDIILECVYCKQQLLRRE**VYDFAFRDL**CI**VY**RDGNPYAVC 70
 DKCLKFYISKISEYRHICYSVYGTTLQOYNKPLCDLLIRCINCQKPLCPEEKQRHLDKKQRFHNIIRGWT 140
 GRMSSCRSSRTRRETQL

SEQ ID NO:13
 >gi|56583270|ref|NP_040979.2| matrix protein 2 [Influenza A virus (A/Puerto
 Rico/8/34 (H1N1))]
 MSLLTEVETPIRNEWGCRCNGSSDPLAIAANIIGILHLILWILDRLFFKCIYRRFKYGLKGGPSTEGVPK
 25 SMREEYRKEQQSAVDADDGHFVSIELE

SEQ ID NO:14
 >gi|8486139|ref|NP_040987.1| PB2 protein [Influenza A virus (A/Puerto
 Rico/8/34 (H1N1))]
 30 MERIKELRNLSQSRTEILTCTVDHMAIIKKYTSGRQEKNPALRMKMMAMKYPITADKRITEMIPER
 NEQGQTLWSKMNDAGSDRMVSPPLAVTWWNRNGPMTNTVHYPKIYKTYFERVERLKHGTFGPVHFRNQVK
 IRRRVDINPGHADLSAKEAQDVIMEVVFNPVEVGARILTSQSQTITKEKKEELQDCKISPLMVAYMLERE
 LVRKTRFLPVAGGTSSVYIEVLHLTQGTCEWQMYTPGGEVKNDDVDQSLIIAARNIVRRAAVSADPLASL
 LEMCHSTQIGGIRMVDILKQNPTEEQAVGICKAAMGLRISSSFSGGFTFKRTSGSSVKREEEVLGTGNLQ
 35 TLKIRVHEGYEEFTMVGRRAAILRKATRRLIQLIVSGRDEQSI AEAIIVAMVFSQEDCMIKAVRGDLNF
 VNRRANQRLNPMHQLLRHFQKDAKVLQNWGVEPIDNVGMIGILPDMTPSIEMSMRGVRI SKMGVDEYSS
 TERVVVSI DRFLVRDQRGNVLLSPEEVSETQGTEKLITITYSSSMWEINGPESVLVNTYQWIIIRNWETV
 KIQWSQNPTMLYNKMEFEPFQSLVPKAI RQYSGFVRTLFQQMRDVLGTFDTAQIIKLLPFAAAPPKQSR
 MQFSSFTVNVRGSGMRILVRGNSPVFNYNKATKRLTVLGKDAGTLTEDPDEGTAGVESAVLRGFLILGKE
 40 DRRYGFALSINELSNLAKGEKANVLIGQGDVVLV MKRKRDSSILTDSTATKRIRMAIN

SEQ ID NO:15
 >gi|8486137|ref|NP_040986.1| polymerase PA [Influenza A virus (A/Puerto
 Rico/8/34 (H1N1))]
 45 MEDFVRQCFNPMIVELA EAKTMKEYGEDLKIETNKFAAICTHLEVCFMYSDFHFINEQGESI IVELGDPNA
 LLKHRFEIIEGRDRTMAWTVVNSICNTTGA EKPKFLPDLYDYKENRFIEIGVTRREVHIYYLEKANKIKS
 EKTHIHIFSFTGEEMATKADYTLDEESRARIKTRLFTIRQEMASRGLWDSFRQSERGEETIEERFEITGT
 MRKLADQSLPPNFSSLENFRAYVDGFEPNGYIEGKLSQMSKEVNARIEPFLKTTPRPLRLPNGPPCSQRS
 KFLMDALKLSIEDPSHEGEGIPLYDAIKCMRTFFGWKEPNVVKPHEKGINPNYLLSWKQVLAELQDIEN
 50 EEKIPKTKNMKKTSQLKWALGENMAPEKVD FDDCKDVGD LKQYDSDEPELRSLSWIQNEFNKACELTDS
 SWIELDEIGEDVAPIEHIA SMRRNYFTSEVSHCRATEYIMKGVYINTALLNASCAAMDDFQ LIPMISKCR
 TKEGRKTNLYGFIIKGRSHLRNDTDVVNFVSM EFSLTDPRLPHKWEKYCVLEIGDMLLRSAIGQVSRP
 MFLYVRTNGTSKIKMKWGMEMRCLLQSLQ QIESMIEAESSVKEKDMTKEFFENKSETWPIGESPKGVEE
 SSIGKVCRTLLAKSVFNSLYASPQLEGFSAESR KLLLI VQALRDNL EPGTFDLGGLYEAIEECLINDPWV
 55 LLNASWFNSFLTHALS

SEQ ID NO:16
 >gi|8486133|ref|NP_040984.1| nonstructural protein NS1 [Influenza A virus
 (A/Puerto Rico/8/34 (H1N1))]

MDPNTVSSSFQVDCFLWHVRKRVDQELGDAPFLDRLRRDQKSLRGRGSTLGLDIETATRAGKQIVERILK
EESDEALKMTMASVPASRYLTDMTLEEMSREWSMLIPKQKVAGPLCIRMDQAIMDKNIILKANFSVIFDR
LETLLILLRAFTEEGAIVGEISPLPSLPGHMTAEDVKNAVGVLIIGGLEWNDNTVRVSETLQRFARSSNENG
RPPLTPKQKREMAGTIRSEV

5

SEQ ID NO:17

>gi|8486132|ref|NP_040983.1| nonstructural protein NS2 [Influenza A virus (A/Puerto Rico/8/34 (H1N1))]

10 MDPNTVSSSFQDILLRMSKMQLESSSEDINGMITQFESLKLYRDSLGEAVMRMGDLHSLQNRNEKWREQLG
QKFEEIRWLIIEVRHKLKVTENSFEQITFMQALHLLLEVEQEIRTFSFQLI

SEQ ID NO:18

>gi|8486128|ref|NP_040981.1| neuraminidase [Influenza A virus (A/Puerto Rico/8/34 (H1N1))]

15 MNPNQKIITIGSICLVGLISLILQIGNIISIWISHSIQTGSQNHTGICNQNIITYKNSTWVKDTTSVIL
TGNSSLCPIRGWAIYSKDNSIRIGSKGDVFEVIREPFISCSHLECRTEFFLTQGALLNDRHSNGTVKDRSPY
RALMSCPVGEAPSPYNSRFESVAWSASACHDGMGWLITIGISGPDNGAVAVLKYNGIITETIKSWRKKILR
TQSEACACVNGSCFTIMTDGPSDGLASYKIFKIEKGKVTKSIELNAPNSHYEECSYPTDGKVMCVCARDN
20 WHGSSNRPWVSFDQNLDYQIGYICSGVFGDNPRPKDGTGSCGPVYVDGANGVKGFYSRYGNGVWIGRTKSH
SSRHGFEMIWDPNWGTETDSKFSVRQDVVAMTDWSGYSGSFVQHPELTGLDCIRPCFWVELIRGRPEKT
IWTSSASSISFCGVNSDITVDWSWPDGAELPFTIDK

SEQ ID NO:19

25 >gi|8486126|ref|NP_040980.1| haemagglutinin [Influenza A virus (A/Puerto Rico/8/34 (H1N1))]

MKANLLVLLCALAAADADTICIGYHANNSTDTVDTVLEKNVTVTHSVNLLLEDHNGKLCRLKGIAPLQLG
KCNIAGWLLGNPECDPLLPVRSWSYIVETPNSENGICYPGDFIDYEELREQLSSVSSFERFEIFPKESSW
PNHNTTKGVTAACSHAGKSSFYRNLLWLTEKEGSSYPKLKNSYVNKKGKEVLVLWGIHHPNSKDDQNIYQ
NENAYVSVVTSNYNRRFTPEIAERP KVRDQAGRMNYYWTLKPGDTIIFEANGNLIAPRYAFALSRGFGS
30 GIITSNASMHECNTKCTPLGAINSSLPFQNIHPVTIGCEPKYVRSALRMVMTGLRNIPSIQSRGLFGAI
AGFIEGGWTGMIDGWYGYHHQNEQGSYAADQKSTQNAINGITNKVNSVIEKMNIQFTAVGKEFNKLEKR
MENLNKKVDDGFLDIWTYNAELLVLLLENERTLDFHDSNVKNLYEKVKSQKNNAKEIGNGCFEFYHKCDN
ECMESVRNGTYDYPKYSEESKLNREKVDGVKLESMGIYQILAIYSTVASSLVLLVSLGAISFWMCNNGSL
QCRICI

35

SEQ ID NO:20

>gi|8486123|ref|NP_040978.1| matrix protein 1 [Influenza A virus (A/Puerto Rico/8/34 (H1N1))]

40 MSLLTEVETYVLSIIPSGPLKAEIAQRLEDVFAGKNTDLEVLMEWLKTRPILSPLTKGILGFVFTLTVP
ERGLQRRRFVQNALNGDPNMDKAVKLYRKLKREITFHGAKEISLSYSAGALASCMGLIYNRMGAVTT
EVAFLVCATCEQIADSQHRSHRQMVTTTNPLIRHENRMVLASTTAKAMEQMAGSSEQAAEAMEVASQAR
QMVQAMRTIGTHPSSSAGLKNDDLLENLQAYQKRMGVQMQRFK

SEQ ID NO:21

45 >gi|83031685|ref|YP_418248.1| PB1-F2 protein [Influenza A virus (A/Puerto Rico/8/34 (H1N1))]

MGQEQDTPWILSTGHISTQKRQDGOQTPKLEHRNSTRLMGHCQKTMNQVVMKQIVYWKQWLSLRNPILV
FLKTRVLKRWRLFSKHE

50

SEQ ID NO:22

>gi|8486135|ref|NP_040985.1| polymerase 1 PB1 [Influenza A virus (A/Puerto Rico/8/34 (H1N1))]

55 MDVNPTLLFLKVPQAIAISTTFPYTGDPYPYSHGTGTGYTMDTVNRTHQYSEKARWTTNTETGAPQLNPID
GPLPEDNEPSGYAQTDCVLEAMAFLEESHPIFENSCTETMEVVQQTRVDKLTQGRQTYDWTNLNRNPAA
TALANTIEVFRSNGLTANESGRILIDFLKDVMESSMKKEEMGITTHFQKRVRVNDMTKKMITQRTIGKRKQ
RLNKRSYLIRALTNTMTKDAERGLKRRAIATPGMQIRGFVYFVETLARSICEKLEQSGLPVGGNEKKA
KLANVVRKMMTNSQDTELSLITITGDNTKWNENQNPRLAMITYMTRNQPEWFRNVLSIAPIMFNSKMAR
LGKGYMFESKMKLRTOIPAEMLASIDLKYFNDSTRKKIEKIRPLLIEGTASLSPGMMGMFMNMLSTVLG
VSILNLGQKRYTKTTYWWDGLQSSDDFALIVNAPNHEGIQAGVDRFYRTCKLHGINMSKKKSYINRTGTF

EFTSFFYRYGFVANFSMELPSFGVSGSNESADMSIGVTVIKNNMINNDLGPATAQMALQLFIKDYRYTYR
 CHRGD^TQIQTRRSFEIKKLWEQTRSKAGLLVSDGGPNLYNIRNLHIPEVCLKWELMDE^DYQGRLCNPLNP
 FVSHKEIESMNNAVMMPAHGPAKNMEYDAVAT^THSWIPKRNRSILNTSQRGVLEDEQMYQRCNLF^EKFF
 PSSSYRRPVGISSMVEAMVSRARIDARIDFESGRIKKEEFTEIMKICSTIEELRRQK

5

SEQ ID NO:23

>gi|8486130|ref|NP_040982.1| nucleocapsid protein [Influenza A virus
 (A/Puerto Rico/8/34 (H1N1))]

MASQGT^KRSYEQMETDGERQ^NATEIRASVGMIGGIGRFYIQMCTELKLS^DYEGRLIQNSLTIERM^VLSA
 10 FDERRNKYLEEHPSAGKDPKKTGGPIYRRVNGKWMRELILYDKEEIRRIWRQANNGDDATAGLTHMMIWH
 SNLNDATYQ^RTRALV^RTGM^DPRMCSLMQGSTLPRRS^GAAGAAVKG^VGT^MVMELVRMIKRGINDRN^FWRGE
 NGRKTRIA^YERM^CNILKGKFQ^TAAQKAMMDQVRES^RDPGNAEFEDLTFLARSALILRGSVAHKSCLPACV
 YGP^AVASGYDFERE^GYSLVGIDPFRLLQNSQVYSLIRPNENPAHKSQ^LVWMACHSAA^FEDLRVLSFIKGT
 KVVPRGKLSTRGVQ^IASNENMETMESSTLELRSRYWAI^RTRSGGNTNQQRASAGQ^ISIQPTFSVQ^RNLPF
 15 DRTTVMAAFTGNT^EGRTSDMRTEIIRMMESARPE^DVSFQGRGVFELSDEKAASPIVPSFDMSNEGSYFFG
 DNAEEYDN

SEQ ID NO:24

20 >gi|73918826|ref|YP_308855.1| polymerase 2 [Influenza A virus
 (A/Korea/426/1968 (H2N2))]

MERIKELRNLMSQ^SRTREILTKTTVDHMAIIKKYTSGRQ^EKNPSLRMKWMMAMKY^PITADKRITEMVPER
 NEQGQTLWSKMSDAGSD^RVMVSP^LAVTWNNRNGPMTSTVHYPK^IYKTYFEKVERLKHGTFGPVHFRNQVK
 IRRRVDINPGHADLSAKEAQDVIMEVVF^PNEVGARILTS^ESQ^LTITKEKKEELQDCKISPLMVAYMLERE
 25 LVRKTRFLPVAGGTSSVYIEVLH^LTQGT^CWEQMYTPGGEVRNDDVDQSLIIAARNIVRRAAVSADPLASL
 LEMCHSTQIGGTRMVDILRQNPTEEQA^VDICKAAMGLRISSSFSGGFTFKRTSGSSI^KREEEVLTGNLQ
 TLKIRVHEGYEEFTMVGKRATAILRKATRRLV^QLIVSGRDEQ^SIAEAIIVAMVFSQEDCMIKAVRGDLN^F
 VNRRANQRLNPMHQLLRHFQKDAKVL^FQNWGIEHIDNVGMIGVLPDMTPSTEMSMRGIRVSKMGVDEYSS
 TERVVVSIDRFLRVRDQ^RGNVLLSP^EEVSETQGT^EKL^TITYSSSMWEINGPESVLVNTYQWII^RNWETV
 30 KIQWSQNPTMLYNKMEFEPFQSLVPKAI^RGQYSGFVRTLFQQMRDVLGT^FDTTQIIKLLPFAAAPPKQSR
 MQFSSLT^VNVRGSGMRILVRGNSPVFNYNK^TTKRLTILGKDAGTLTEDPDEGTSGVESAVLRGFLILGKE
 DRRYGPALSINELSTLAKGEKANVLIGQGDVVLV^MKRKR^DSSIL^TSDSQ^TATKRI^RMAIN

35

SEQ ID NO:25

>gi|73919145|ref|YP_308850.1| hemagglutinin [Influenza A virus
 (A/Korea/426/68 (H2N2))]

MAIIYLILLFTAVRGDQICIGYHANNSTEKVD^TILERNVT^VTHAKDILEKTHNGKLCKLNGIPPLELGDC
 40 SIAGWLLGNPECDRLLSVPEWSYIMEKENPRYSLCYPGSFNDYEELKHL^LSSVKHFEKVKILPKDRWTQH
 TTTGGSWACAVSGKPSFFRN^MVWLTRKGSNYPVAKGSYNNTSGEQMLIIWG^VHHPNDEAEQ^RALYQ^NVT
 YVS^VATSTLYKRSIPEIAARPKV^NGLGRMEFSWTL^LDMWD^TINFESTGNLVAPEYGFKISKRGSSGIMK
 TEGTLENCETK^CQTPLGAINTTLPFHN^VHPLTIGEC^PKYVKSEKLVLATGLRNVPQIESRGLFGAIAGFI
 EGGWQGMVDGWYGYHHSNDQGS^GYAADKESTQKAFNGITNKVNSVIEKMNTQFEAVGKEFSNLEKRL^NL
 45 NKKMEDGFLDVW^TYNAELLVLMENERTLDFHDSNVKNLYDKVRMQLRDNV^KELGNGCFEFYHKCDNECMD
 SVKNGTYDYPKYEEESKLNRNEIKGVKLSSMGVYQILAIYATVAGSLSLA^IMMAGISFWMC^SNGSLQCRIC
 I

SEQ ID NO:26

50 >gi|73912688|ref|YP_308854.1| membrane protein M1 [Influenza A virus
 (A/Korea/426/68 (H2N2))]

MSLLTEVETYVLSIVPSGPLKAEIAQ^RLEDV^FAGKNTDLEALMEWLKTRPILSPLTKGILGFVFTLTVP^S
 ERGLQRRRFVQ^NALNGNDPNMDRAVKLYRKLKREITFHGAKEVALSYSAGALASCMGLIYNRMGA^VTT
 EVAFAVVCATCEQ^IADSQHRSHRQ^MVTTTNPLIRHENRMVLASTTAKAMEQMAGSSEQA^EAAMEVASQ^AR
 55 QMVQAMRAIGTPSSSAGLKDDLLENLQAYQKRMGVQMQR^FK

SEQ ID NO:27

>gi|73912687|ref|YP_308853.1| membrane protein M2 [Influenza A virus
 (A/Korea/426/68 (H2N2))]

MSLLTEVETPIRNEWGCRCNDSSDPLVVAASIIGILHFILWILDRLFFKCIYRFFKHGLKRGPPSTEGVPE
SMREEYRKEQQSAVDADDSHFVSIELE

SEQ ID NO:28

- 5 >gi|73912685|ref|YP_308852.1| polymerase PA [Influenza A virus
(A/Korea/426/68 (H2N2))]
MEDFVRQCFNPMIVELAEKAMKEYGEDLKIETNKFAAICTHLEVCFMYSDFHFINEQGESIMVELDDPNA
LLKHRFEIIEGRDRTMAWTVVNSICNTTGAEKPKFLPDLYDYKENRFIEIGVTRREVHIYYLEKANKIKS
ENTHIHIFSFTGEEMATKADYTLDEESRARIKTRFLTIRQEMANRGLWDSFRQSERGEETIEERFEITGT
10 MRRLADQSLPPNFSCLENFRAYVDGFEPNGYIEGKLSQMSKEVNAKIEPFLKTPRPRLPDGPPCFQRS
KFLMDALKLSIEDPSHEGEGIPLYDAIKCMRTFFGWKEPYIVKPHEKGINPNYLLSWKQVLAELQDIEN
EEKIPRTKNMKKTSQKWLALGENMAPEKVDFDNCRDISDLKQYDSDEPELRSLSWQNEFNKACELTDS
IWIELDEIGEDVAPIEHASMRNYFTAEVSHCRATEYIMKGVYINTALLNASCAAMDDFQLIPMISKCR
TKEGRKTNLYGFIIKGRSHLRNDTDVNVFVSMFSLTDPRLPHKWEKYCVLEIGDMLLRSAIGQMSRP
15 MFLYVRTNGTSKIKMKWGMEMRPCLLQSLQQIESMVEAESSVKEKDMTKEFFENKSETWPIGESPKGVVEE
GSIGKVCRTLLAKSVFNSLYASPQLEGFSAESRKLILLVQALRDNLPGTDFDLGGLYEAEELCLNDPWV
LLNASWFNSFLTHALR

SEQ ID NO:29

- 20 >gi|73921833|ref|YP_308877.1| PB1-F2 protein [Influenza A virus
(A/Korea/426/68 (H2N2))]
MGQEQDTPWTQSTEHINIQRGSGQQTRKLERPNLTQLMDHYLRTMNQVDMHKQTASWKQWLSLRNHTQE
SLKIRVLKRWKLFNKQEWTN

SEQ ID NO:30

- 25 >gi|73912683|ref|YP_308851.1| PB1 polymerase subunit [Influenza A virus
(A/Korea/426/68 (H2N2))]
MDVNPTELLFLKVPQAQNAISTTFPYTGDPYPYSHGTGTGYTMDTVNRTHQYSEKKGWTTNTTETGAPQLNPID
GPLPEDNEPSGYAQTDCVLEAMAFLEESHPIGIFENSCLTMEVIQQTRVDKLTQGRQTYDWTNLNRNQPA
30 TALANTIEVFRSNGLTANESGRLIDFLKDVIESMDKEEMEITTHFQRKRVRDNTKKMVTQRTIGKKKQ
RLNKRSYLIRALTNTMTKDAERGKLRRAIATPGMQIRGFVHFVETLARNICEKLEQSGLPVGGNEKKA
KLANVVRKMTNSQDTELSFTITGDNTKWNENQNPRVFLAMITYITRNQPEWFRNVLSIAPIMFSNKMAR
LGKGYMFESKSMKLRTQIPAEMLASIDLKYFNSTRKKIEKIRPLLDGTVSLSPGMMGMFMNMLSTVLG
VSIILNLGQKKYTKTTYWWDGLQSSDDFALIVNAPNHEGIQAGVNRFYRTCKLVGINMSKKKSYINRTGTF
35 EFTSFFYRYGFVANFSMELPSFGVSGINESADMSIGVTVIKNMINNDLGPATAQMALQLFIKDYRYTYR
CHRGDTQIQTRRSFELKKLWEQTRSKAGLLVSDGGSNLYNIRNLHIPEVCLKWELMDEYQGRLCNPLNP
FVSHKEIESVNNAVMPAHGPAKSMEYDAVATTHSWTPKRNRSILNTSQRGILEDEQMYQKCCNLFEKFF
PSSSYRRPVGISSMVEAMVSRARIDARIDFESGRIKKEEFAEIMKICSTIEELRRQK

SEQ ID NO:31

- 40 >gi|73921567|ref|YP_308869.1| non-structural protein NS2 [Influenza A virus
(A/Korea/426/68 (H2N2))]
MDSNTVSSFQDILLRMSKMQLGSSSEDNLGMITQFESLKYRDSLGEAVMRMGDLHSLQNRNGKWREQLG
QKFEEIRWLIEEVRHRLKITENSFEQITFMQALQLLFEVEQEIRTFQSLI

SEQ ID NO:32

- 45 >gi|73921566|ref|YP_308870.1| non-structural protein NS1 [Influenza A virus
(A/Korea/426/68 (H2N2))]
MDSNTVSSFQVDCFLWHVRKQVVDQELGDAPFLDRLRRDQKSLRGRGSTLDLDIEAATRVGKQIVERILK
50 EESDEALKMTMASAPASRYLTDMTIEELSRDWFMLMPKQKVEGPLCIRIDQAIMDKNIMKANFSVIFDR
LETLLILLRAFTEEGAIVGEISPLPSLPGHITIEDVKNAIGVLIGGLEWNDNTVRVSKTLQRFAWRSSNENG
RPPLTPKQKRKMARTIRSKVRRDKMAD

SEQ ID NO:33

- 55 >gi|73921307|ref|YP_308871.1| nucleoprotein [Influenza A virus
(A/Korea/426/68 (H2N2))]
MASQGTKRSEYQMETDGERQNATEIRASVGKMGIDGIGRFYIQMCTELKLSDYEGRLIQNSLTIERMVLSA
FDERNKYLEEHPSAGKDPKKTGGPIYKRVGKWMRELVLVDKEEIRRIWRQANNQDDATAGLTHMMIWH
SNLNDTTYQRTALVRTGMDPRMCSLMQGSTLPRRSGAAGAAVKGVGTMVMEIIRMIKRGINDRNFWRGE

- NGRKTRSAYERMCNILKGKFQTAAQRAMMDQVRESRNPNGNAEIEDLIFLARSALILRGSAVHKSCLPACV
YGPALASGYNFEKEGYSLVGIDPFKLLQNSQVYSLIRPNENPAHKSQLVWVWMACNSAAAFEDLRVLSFIRGT
KVS PRGKLSTRGVQIASNENMDTMESSSTLELRSRYWAI RTRSGGNTNQQRASAGQISVQPAFSVQRNLFP
DKPTIMAAFTGNTEGRTSDMRAEII RMMEGAKPEEMS FQGRGVFELSDEKATNPVPSFDMSEGSYFFG
5 DNAEEYDN
- SEQ ID NO:34
>gi|73921304|ref|YP_308872.1| neuraminidase [Influenza A virus
(A/Korea/426/68 (H2N2))]
10 MNPNQKIITIGSVSLTIATVCFLMQIAILVTTVTTLHFKQHECDSPASNQVMPCEPIIIERNITEIVYLN
TTIEKEICPEVVEYRNWSKPQCQITGFAPFSKDNSIRLSAGGDIWVTREPYVSCDPGKCYQFALGQGTTL
DNKHSNDTIHDRI PHRTLLMNELGVPFHLGTRQVCVAVSSSSCHDGKAWLHVCTGDDKNATASFIYDGR
LMDSIGSWSQNILRTQESECVCINGTCTVVMTDGSASGRADTRILFIEEGKIVHISPLSGSAQHVEECSC
YPRYPDVRCICRDNWKGSNRPVIDINMEDYSIDSSYVCSGLVGDTPRNDRSSNSNCRNPNNERNPGVK
15 GWAFDNGDDVVMGRTISKDLRSYETFKVIGGWSTPNSKSQINRQVIVDSNNWSGYSGIFSVGEKRCINR
CFYVELIRGRQOQETRVWWTNSIVVFCGTS GTYGTGSWPDGANINFMPI
- SEQ ID NO:35
20 >gi|73919213|ref|YP_308844.1| nonstructural protein 2 [Influenza A virus
(A/New York/392/2004 (H3N2))]
MDSNTVSSFQDILLRMSKMQLGSSSEDNLNGMITQFESLKIYRDSLGEAVMRMGDLHLLQNRNGKWREQLG
QKFEEIRWLIEEVRHRLKTTENSFEQITFMQALQLLFEVEQEIRTFSFQLI
- SEQ ID NO:36
25 >gi|73919212|ref|YP_308845.1| nonstructural protein 1 [Influenza A virus
(A/New York/392/2004 (H3N2))]
MDSNTVSSFQVDCFLWHIRKQVVDQELSDAPFLDRLRRDQRSIRGRGNTLGLDIKAATHVGKQIVEKILK
EESDEALKMTMVSTPASRYITDMTIEELSRNWFMMLMPKQKVEGPLCIRMDQAIMEKNIMLKANFSVIFDR
30 LETIVLLRAFTEEGAIVGEISPLPSFPGHGTIEDVKNAIGVLIGGLEWNDNTVRVSKNLQRFARSSNENG
GPPLTPKQKRKMARTARSKV
- SEQ ID NO:37
35 >gi|73919207|ref|YP_308839.1| hemagglutinin [Influenza A virus (A/New
York/392/2004 (H3N2))]
MKTIIALS YILCLVFAQKLP GNDNSTATLCLGHHAVPNGTIVKTITNDQIEVTNATELVQSSSTGGICDS
PHQILDGENCTLIDALLGDPQCDGFQNKWDLFVERSKAYSNCYPYDVPDYASLSLVASSGTLEFNES
FNWTVGTQNGTSSACKRRSNNSSFFSRLNWLTHLKFKYPALNVTMPNNEKFDKLYIWGVHHPGTDNDQISL
YAQASGRITVSTKRSQQTVIPSIGSRPRI RDVP SRISIYWTIVKPGDILLINSTGNLIAPRGYFKIRSGK
40 SSIMRSDAPIGKCNSECITPNGSIPNDKPFQNVNRITYGACPRYVKQNTLKLATGMRNVPEKQTRGIFGA
IAGFIENGWEGMVDGWYGFRHQNSEGTGQAADLKSTQAAINQINGKLNRLIGKTNEKFHQIEKEFSEVEG
RIQDLEKYVEDTKIDLWSYNAELLVALENQHTIDLTDSEMKNL FERTKKQLRENAEDMGNGCFKIYHKCD
NACIGSIRNGTYDHDVYRDEALNNRFQIKGVELKSGYKDWILWISFAISCFLLCVALLGFIMWACQKGN
45 RCNICI
- SEQ ID NO:38
50 >gi|73919153|ref|YP_308840.1| matrix protein 2 [Influenza A virus (A/New
York/392/2004 (H3N2))]
MSLLTEVETPIRNEWGCRCNDSSDPLVVAASIIGILHLILWILDRLFFKCVYRLFHKHGLKRG PSTEGVPE
SMREEYRKEQQNAVDADDSHFVSIELE
- SEQ ID NO:39
55 >gi|73919152|ref|YP_308841.1| matrix protein 1 [Influenza A virus (A/New
York/392/2004 (H3N2))]
MSLLTEVETYVLSIVPSGPLKAEIAQRLEDVFAGKNTDLEALMEWLKTRPILSPLTKGILGFVFTLTVP
ERGLQRRRFVQNALNGNDPNNMDKAVKLYRKLKREITFHGAKEIALSYAGALASCMGLIYNRMGAVTT
EVAFGILVCATCEQIADSQHRSHRQM VATTNPLIKHENRMVLASTTAKAMEQMAGSSEQAAEAMEIASQAR
QMVQAMRAVGTHPSSSTGLRDDLLENLQTYQKRMGVQMQRFK

SEQ ID NO:40

>gi|73919150|ref|YP_308848.1| PB1-F2 protein [Influenza A virus (A/New York/392/2004 (H3N2))]
MEQEQTDPWTQSTEHTNIQRRGSGRQIQKLGHPNSTQLMDHYLRIMSQVDMHKQTVSWRLWPSLKNPTQV
5 SLRTHALKQWKSFNKQGWTN

SEQ ID NO:41

>gi|73919149|ref|YP_308847.1| polymerase PB1 [Influenza A virus (A/New York/392/2004 (H3N2))]
10 MDVNPTLLFLKVPAQNAISTTFPYTGDPYPYSHGTGTGYTMDTVNRTHQYSEKKGWTTNTETGAPQLNPID
GPLPEDNEPSGYAQTDCVLEAMAFLEESHPIGIFENSCLTMEVVQQTRVDKLTQGRQTYDWTNLNRNQPA
TALANTIEVFRSNGLTANESGRLIDFLKDVMEESMDKEEMEITTHFQRKRVRDNMTKKMVTQRTIGKKKQ
RVNKRGYLIRALTTLNMTKDAERGKLRRAIATPGMQIRGFVYFVETLARSICEKLEQSGLPVGGNEKKA
15 KLANVVRKMMTNSQDTELSFTITGDNTKWNENQNPRMFLAMITYITKNQPEWFRNILSIAPIMFSNKMAR
LGKGYMFESKRMKLRTQIPAEMLASIDLKYFNSTRKKIEKIRPLIDGTASLSPGMMMGFMNMLSTVLG
VSVNLNGQKKYTKTTYWWDGLQSSDDFALIVNAPNHEGIQAGVDRFYRTCKLVGINMSKKKSYINKTGTF
EFTSFFYRYGFVANFSMELPSFGVSGINESADMSIGVTVIKNMINNDLGPATAQMALQLFIKDYRYTYR
CHRGDTQIQTRRSFELKKLWDQTQSRAGLLVSDGGPNLYNIRNLHIPEVCLKWELMDENYRGRLCNPLNP
20 FVSHKEIESVNNNAVMPAHGPAKSMEDAVATTHSWNPKRNRSLNTSQRGILEDEQMYQKCCNLFEKFF
PSSSYRRPIGISSMVEAMVSRARIDARIDFESGRIKKEEFSEIMKICSTIEELRRQK

SEQ ID NO:42

>gi|73919147|ref|YP_308843.1| nucleocapsid protein [Influenza A virus (A/New York/392/2004 (H3N2))]
25 MASQGTKRSEYQMETDGRQNAIEIRASVGKMGIDGIGRFYIQMCTELKLSDEHGRLIQNSLTIEKMLVLSA
FDERRNKYLEEHPSAGKDPKKTGGPIYRRVDGKWMRELVLVDKEEIRRIWRQANNGEDATAGLTHIMIWH
SNLNDATYQRTALVRTGMDPRMCSLMQGSTLPRRSGAAGAAVKGIGTMVMEIIRMVKGINDRNFWRGE
NGRKRTRSAYERMCNLIKGFQTAQRAMVDQVRESRNPNAEIEDLI FLARSALILRGSAVHKSCLPACA
YGPVASSGYDFEKEGYSLVGIDPFKLLQNSQIYSLIRPNENPAHKSQVLVMMACHSAAFEDLRLLSFIRGT
30 KVSPRGKLSTRGVQIASNENMDNMGSSSTLELRSGYWAIRTRSGGNTNQQRASAGQTSVQPTFSVQRNLFP
EKSTIMAAFTGNTEGRTSDMRAEII RMMEGAKPEEVSFRGRGVFELSDEKATNPVPSFDMSEGSYFFG
DNAEEYDN

SEQ ID NO:43

35 >gi|73919136|ref|YP_308842.1| neuraminidase [Influenza A virus (A/New York/392/2004 (H3N2))]
MNPNQKIITIGSVSLTISTICFFMQIAILITTVTLHFQKQYEFNSPPNNQVMLCEPTIIERNITEIVYLTN
TTIEKEMCPKLAEYRNWSKPQCDITGFAPFSKDNSIRLSAGGDIWVTREPYVSCDPDKCYQFALGQGTTL
NNVHSNDTVHDRTPYRTLLMNELGVFPFHLGTQKQVCIWSSSSCHDGKAWLHVCTGDDKNATASFIYNGR
40 LVDSIVSWSKKILRTQESECVCINGTCTVVMTDGASGKADTKILFIEEGKIIHTSTLSGSAQHVEECSC
YPRYPGVRVCVRDNWKGSNRPVVDINIKDYSIVSSYVCSGLVGDTPRKNDSSSSSHCLDPNNEEGGHGVK
GWAFDDGNDVMMGRTISEKLRSYETFKVIEGWSPKNSKLQINRQVIVDRGNRSGYSGIFSVEGKSCINR
CFYVELIRGRKEETEVLWTSNSIVVFCGTSPTYGTGSWPDGADINLMPI

45 SEQ ID NO:44

>gi|73919134|ref|YP_308846.1| polymerase PA [Influenza A virus (A/New York/392/2004 (H3N2))]
MEDFVRQCFNPMIVELAEKAMKEYGEDLKIETNKFAAICTHLEVCFMYSDFHFINEQGESIVVELDDPNA
LLKHFREIIEGRDRTMAWTVNSICNTTGAEKPKFLPDLYDYKENRFIEIGVTRREVHIYYLEKANKIKS
50 ENTHIHIFSFTGEEIATKADYTLDEESRARIKTRLFTIRQEMANRGLWDSFRQSERGEETIEEKFEISGT
MRRADQSLPPKFSCLENFRAYVDGFEPNGCIEGKLSQMSKEVNAKIEPFLKTTTPRIKLPNGPPCYQRS
KFLMLDALKLSIEDPSHEGEGIPLYDAIKCIKTFFGWKEPYIVKPHEKGINSNYLLSWKQVLSELQDIEN
EEKIPRTKNMKKTSQKWLWALGENMAPEKVDVDFNCRDISDLKQYDSDEPELRSLSWQNEFNKACELTDS
IWIELDEIGEDVAPIEYIASMRNYFTAEVSHCRATEYIMKGVYINTALLNASCAAMDDFQLIPMISKCR
55 TKEGRRKTNLYGFIIKGRSHLRNDTDVVNFVSMFSLTDPRLPHKWEKYCVLEIGDMLLRSAIGQISRP
MFLYVRTNGTSKVKMKWGMEMRCLLQSLQQIESMIEAESSIKEKDMTKEFFENKSEAWPIGESPKGVEE
GSIGKVCRTLLAKSVFNSLYASPOLEGFSAESRKLILLVQALRDNLPGTDFDLGGLYEAEIEELINDPWV
LLNASWFNSFLTHALK

SEQ ID NO:45

>gi|73919060|ref|YP_308849.1| polymerase PB2 [Influenza A virus (A/New York/392/2004 (H3N2))]

5 MERIKELRNLMSSQSRTEILTKTTVDHMAIIKKYTSGRQEKNP SLRMKMMAMKYPITADKRITEMVPER
NEQGQTLWSKMSDAGSDRVMVSP LAVTWNNRNGPVASTVHYPKVYKTYFDKVERLKHGTFGPVHFRNQVK
IRRRVDINPGHADLSAKEAQDVIMEVVPNEVGARILTS ESQLTITKEKKEELRDCKISPLMVAYMLERE
LVRKTRFLPVAGGTSSIIYIEVLHLTQGT CWEQMYTPGGEVRNDDVDQSLIIAARNIVRRAAVSADPLASL
LEMCHSTQIGGTRMVDILRQNPTEEQAVDICKAAMGLRISSSFSGGFTFKRTSGSSVKKEEEVLTGNLQ
10 TLKIRVHEGYEEFTMVGKRATAILRKATRRLVQLIVSGRDEQSI AEAIIVAMVFSQEDCMIKAVRGDLNF
VNRRANQRLNPMHQLLRH FQKDAKVL FQNWGIEHIDSVGMGVGLPDMTPSTEMSMRGIRVSKMGVDEYSS
TERVVVSI DRFLRVRDQ RGNVLLSPEEVSETQGT ERLTITYSSSMWEINGPESVLVNTYQWII RNWEAV
KIQWSQNPAMLYNKMEFEPFQSLVPKAIRSQYSGFVRTLFQQMRDVLGTFD TTQIIKLLPFAAAPPKQSR
MQFSSSLTVNVRGSGMRILVRGNSPVFNYNKTTKRLTILGKDAGTLIEDPDESTSGVESAVLRGFLIIGKE
15 DRRYGPALSINELSNLAKGEKANVLI GQGDVVLVMKRKRDS SILTDSQTATKRIRMAIN

SEQ ID NO:46: CMV Protein IE122:

20 >gi|39841910|gb|AAR31478.1| UL122 [Human herpesvirus 5]
MESSAKRKMDPDNPDEGPSSKVP RPETPVTKATTF LQTMLRKEVNSQLSLGDP LFPPELAEESLKTFEQVT
EDCNENPEKDVLAELGDI LAQAVNHAGIDSSSTGHTLTTHSCSVSSAPLNKPTPTSVAVINTPLPGASAT
PELSPRKKPRKTTRPFKVIIKPPVPPAPIMLPLIKQEDIKPEPDFTIQYRNKIIDTAGCIVISDSEEEQG
EEVETR GATASSPSTGSGT PRVTSPTHPLSQMNHPLPLDPLARPD EDDSSSSSSSSC SSASDSESESEEMK
25 CSSGGGASVTSSHHGRGGFGSAASSLLSCGHQSSGGASTGPRKKKSKRI SELDNEKVRNIMKDKNTPFCTPNVQTRRG
RVKIDEVSRMFRNTNRSLEYKNLPFTIPSMHQVLDEAIKACKTMQVNNKGIQIIYTRNHEVKSEVD A VRCRLGTMCNLA
LSTPFLMEHTMPVTHPPEVAQRTADACNEG VKAAWSL KELHTHQLCPRSSDYRNMI IHAATFVDLLGALNLC LPLMQKF
PKQVMVRI FSTNQGGFMLPIYETA AKAYAVGQFEQPTETPPEDLD TSLAIEAAIQDLRNKSQ

30 SEQ ID NO:126:

>gi|4927721|gb|AAD33253.1|AF125673_2 E7 [Human papillomavirus type 16]
MHGDTPTLHEYMLDLQPETTDLYCYEQLNDSSEEDEIDGPAGQAEPDRAHYNI VTFCKCDSTLR LCVQ
STHVDIRTLEDLLMGTLGIVCPICSQKP

35

EXAMPLE 1

- 5 The peptides according to the invention used in the following examples were synthesized by Schafer-N as c-terminal amides using the Fmoc-strategy of Sheppard, (1978) J.Chem.Soc., Chem. Commun., 539.

Cell penetration assay

- 10 A set of peptides were biotinylated on N-terminal, and different combinations of aminoacids, with respect to length and type, were added to the sequence box X¹, X³ and X⁴ in the peptides as illustrated by the diagram below. The peptides were tested on cells grown from one individual blood donor.

- 15 Schematic diagram of amino acid sequence of the peptides according to the invention (Each X defines a sequence of amino acids):

X ¹	X ²	X ³	X ⁴	X ⁵
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SP_2 to SP_17 are variants of 51n_Biotin, from a specific native domain on the HCV core protein.

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Intracellular staining for biotinylated peptides

- 96-well U-bottom polystyrene plates (NUNC, cat no: 163320) were used for staining of human PBMCs. Briefly, 8ul of N- or C-terminally biotinylated peptides according to table 1 or table 2 (i.e. 5mM, 2.5mM & 1.25mM tested for each peptide) were incubated at 37°C for 2h
 25 with 40ul of PBMC (12.5 x 10⁶ cells/ml) from blood donors. Cells were then washed 3x with 150ul of Cellwash (BD, cat no: 349524), followed by resuspension of each cell pellet with 100ul of Trypsin-EDTA (Sigma, cat no: T4424), then incubated at 37°C for 5 min. Trypsinated cells were then washed 3x with 150ul of Cellwash (BD, cat no: 349524), followed by resuspension with BD Cytofix/Cytoperm™ plus (BD, cat no: 554715), then incubated at
 30 4°C for 20 min according to manufacturer. Cells were then washed 2x with 150ul PermWash (BD, cat no: 554715). Cells were then stained with Streptavidin-APC (BD, cat no: 554067) & Anti-hCD11c (eBioscience, cat no: 12-0116) according to manufacturer at 4°C for 30 min

aiming to visualize biotinylated peptides & dendritic cells, respectively. Cells were then washed 3x with 150ul PermWash, followed by resuspension in staining buffer (BD, cat no: 554656) before flow cytometry. Dendritic cells were gated as CD11c+ events outside lymphocyte region (i.e. higher FSC & SSC signals than lymphocytes). 200 000 total cells were acquired on a FACSCanto II flow cytometer with HTS loader, and histograms for both total cells & dendritic cells with respect to peptide-fluorescence (i.e. GeoMean) were prepared.

Extracellular staining for biotinylated peptides

96-well U-bottom polystyrene plates (NUNC, cat no: 163320) were used for staining of human PBMCs. Briefly, 8ul of N- or C-terminally biotinylated peptides according to table 1 or table 2 (i.e. 5mM, 2.5mM & 1.25mM tested for each peptide; all peptides manufactured by Schafer) were incubated at 37°C for 2h with 40ul of PBMC (12.5 x 10⁶ cells/ml) from blood donors. Cells were then washed 3x with 150ul of Cellwash (BD, cat no: 349524), then stained with Streptavidin-APC (BD, cat no: 554067) & Anti-hCD11c (eBioscience, cat no: 12-0116) according to manufacturer at 4°C for 30 min aiming to visualize biotinylated peptides & dendritic cells, respectively. Cells were then washed 3x with 150ul of Cellwash (BD, cat no: 349524), followed by resuspension in staining buffer (BD, cat no: 554656) before flow cytometry. Dendritic cells were gated as CD11c+ events outside lymphocyte region (i.e. higher FSC & SSC signals than lymphocytes). 200 000 total cells were acquired on a FACSCanto II flow cytometer with HTS loader, and histograms for both total cells & dendritic cells with respect to peptide-fluorescence (i.e. GeoMean) were prepared.

It was clearly seen that the arginine added on the beginning, middle and in the end improves the ability to enter the cell. It was observed that the variations of arginine decides the ability to enter the cell, best result are achieved with two in the beginning and two in the middle – but variations of different amount of arginines all show good result.

The data shown in the tables below are geomean-value of each tested peptide, as calculated by the FACS Duva software. The Geomean values by trypsinating/Cytofix/Cytoperm:

Table 3

With Cytofix/Cytoperm and trypsination

BC20	Concentration			
	833µM	417 µM	208 µM	104 µM
51n_Biotin	ND	ND	ND	ND
SP 2	3136	5486		1211

SP 3	40456	25905	10788	4308
Sp 4	41491	28008	7540	2946
Sp 5	40534	46060	19065	6152
Sp 6	32577	22195	6514	1850
Sp 7	47922	30503	10481	2766
SP 8	31976	28606	9617	2917
SP 9	11344	6060	2945	1268
SP 10	43389	18607	7684	3709
SP 11	8262	3562	1277	601
SP 12	11089	10869	8679	4316
SP 13	568	394	332	
SP 14	1059	682	392	62
SP 15	1810	477	848	600
SP 16	4558	2240	1260	505
SP 17	30248	29967	12244	1500
SP 18	4869	1334	1971	1038
SP 19	17018	7201	3466	1625
SP 20	7319	2457	1268	445
- ve	0	0	0	0
+ ve	18694	26046	15340	7182
No peptide	224	106	78	168

As shown in table 4 the peptides according to the invention had the ability to allocate through the cell membrane.

- 5 Table 4: Results on DC-cells with low concentration of peptides (208 μ M). The results are sorted in descending order (with respect to BC 16).

Peptide	BC 16	BC43
no pept	4473	2488
NO pept	4003	2359
NO pept	7799	
P-biotin	22058	33545
P-biotin	5568	26212
<i>P-biotin</i>	28169	
n-biotin	3745	1967
N-biotin	3591	3631

N-biotin	2271	
SP61_2	128330	150491
42	74920	65811
SP42_1	73973	
SP17	67428	
SP51_1*	66582	
BI10026	65196	43536
SP11b_1	64047	
42b	61968	19181
SP22	61597	18624
51 biotin	59789	33935
SP12	59274	31222
BI100260	47846	
SP61_3	47182	36803
V10	45430	
120b	44185	
100-22b	43537	28780
BI100260b	37702	
SP12_c	37100	43386
18b	29740	33597
SP13	28539	9129
BI10024b	26892	13002
SP21	26525	4294
61b biotin	25611	24416
SP61b_2	25489	
310-11-n-sc	25236	
100-12	24792	45622
SP5	21358	29146
SP4_c	20673	30786
SP7_c	20041	11832
SP20	19051	3383
190n	18616	23383
SP3_c	18505	26308
SP24	18181	63639
SP6_c	17964	38487
SP42b_1	17628	
SP23	17084	12801
SP5_c	16908	24403
SP7	16064	21414
SP8	15379	17836
BI-050sc5	13648	15549
190	12906	
SP61_4	12829	
190b	12391	20444

SP17_c	11830	
SP25	11552	
SP4	10751	17372
SP6	10233	14468
SP3	9480	10573
SP1_c	9234	
SP8_c	9159	23763
SP10	8627	9216
19	8417	6372
SP9	8356	10084
N13	7982	5953
51b	7000	6082
SP10_c	6904	
SP2	6677	6112
310-11	6537	
SP19	6522	6536
SP11	6490	13230
SP9_c	6180	7758
BI-050sc1	6056	6160
BI-050sc2	6019	11584
SP16	5730	3656
51n	5551	7721
SP18	5545	7751
310-11n	5436	
SP11_c	5371	7878
SP15	5076	2539
N10	4597	
SP14	3704	1629
SP2_c	323	13341
SP13_c		60307
42n		

Example 2

Positive CTL response may alternatively be assayed by ELISPOT assay.

Human IFN-gamma cytotoxic T-cell (CTL) response by ELISPOT assay

- Briefly, at day 1, PBMC samples from HCV patients were incubated in flasks (430 000 PBMCs/cm²) for 2h at 37°C, 5% CO₂ in covering amount of culture media (RPMI 1640 Fisher Scientific; Cat No. PAAE15-039 supplemented with L- Glutamine, (MedProbe Cat. No. 13E17-605E, 10% Foetal Bovine serum (FBS), Fisher Scientific Cat. No. A15-101) and

Penicillin/Streptomycin, (Fisher Scientific Cat. No. P11-010) in order to allow adherence of monocytes. Non-adherent cells were isolated, washed, and frozen in 10% V/V DMSO in FBS until further usage. Adherent cells were carefully washed with culture media, followed by incubation at 37°C until day 3 in culture media containing 2 µg/ml final concentration of hrGM-CSF (Xiamen amoytop biotech co, cat no: 3004.9090.90) & 1 µg/ml hrIL-4 (Invitrogen, Cat no: PHC0043), and this procedure was then repeated at day 6. At day 7, cultured dendritic cells (5 000-10 000 per well) were added to ELISPOT (Millipore multiscreen HTS) plates coated with 0.5 µg/well anti-human γ Interferon together with thawed autologous non-adherent cells (200 000 per well), antigen samples (1-8 µg/ml final concentration for peptide antigens; 5 µg/ml final concentration for Concanavalin A (Sigma, Cat no: C7275) or PHA (Sigma, Cat no: L2769)) & anti-Interferon antibodies (0.03-0.05 µg/ml final concentration for both anti-PD-1 (eBioscience, cat no: 16-9989-82) & anti-PD-L1 (eBioscience, cat no: 16-5983-82)). Plates were incubated overnight and spots were developed according to manufacturer. Spots were read on ELISPOT reader (CTL-ImmunoSpot® S5 UV Analyzer).

Table 5: ELISPOT of core proteins

		Patient A		Patient B		Patient C		Patient D	
	ConA	5090	5035	2605	2725	1355	700	529	512
	No Pep	20	25	10	25	10	0	4	1
	Irr. peptide	10	30	40	5	45	0	1	2
RRSRNLGKVIDTLTCBRLMGYIPLVGA	310-42b	90	115	60	80	70	25	13	11
RRIRNLGRVIETLTGBRLZGYIPLIGA	310-42	75	80	95	30	25	55	13	11
RRGYIPLVGAPLGBRVARALAHGVRV	310-51b	45	40	50	85	45	75	9	13
RRGYLPAVGAPIGBRVIRVIAHGLRL	310-51	40	75	50	60	100	35	9	9
RRNYATGNLPGRRGCSFSIFLLAL	310-61b	45	35	25	45	55	40	15	17
RRGGGQIIGGNYLIPRBGPBIGVRATB	310-11	ND	ND	ND	ND	ND	ND	11	14

Table 5 – Results from IFN- γ ELISPOT run on scaffold peptides with domains derived from HCV proteins

The peptides had in general a higher amount of T-cell spots compared to No Peptide and the irrelevant control peptide. This clearly indicates that the peptides have a positive immunological effect in HCV-patients.

Example 3

The REVEAL & ProVE® Rapid Epitope Discovery System in Detail

Binding properties to HLA for the ninemers listed in table 6 were tested for the following HLA-classes:

HLA-A1, HLA-A2, HLA-A3, HLA-A11, HLA-A24, HLA-A29, HLA-B7, HLA-B8, HLA-B14, HLA-B15, HLA-B27, HLA-B35, HLA-B40.

- 5 The peptides were synthesized as a Prospector PEPscreen®: Custom Peptide Library. Peptides 8-15 amino acids in length were synthesized in 0.5-2mg quantities with high average purity. Quality control by MALDI-TOF Mass Spectrometry was carried out on 100% of samples.
- 10 The REVEAL™ binding assay determined the ability of each candidate peptide to bind to one or more MHC class I alleles and stabilizing the MHC-peptide complex. By comparing the binding to that of high and intermediate affinity T cell epitopes, the most likely immunogenic peptides in a protein sequence can be identified. Detection is based on the presence or absence of the native conformation of the MHC-peptide complex.
- 15 Each peptide is given a score relative to the positive control peptide, which is a known T cell epitope. The score of the test peptide is reported quantitatively as a percentage of the signal generated by the positive control peptide, and the peptide is indicated as having a putative pass or fail result. Assay performance is confirmed by including an intermediate control peptide that is known to bind with weaker affinity to the allele under investigation.

20

Table 6:

HLA-type	A1	A2	A3	A11	A24	A29	B7	B8	B14	B15	B27	B35	B4
Positive Control	100,00	100,00	100,00	100,00	100,00	100,00	100,00	100,00	100,00	100,00	100,00	100,00	100
Intermediate Control	8,41	8,79	10	14,95	10	30,52	12,00	56,60	24,21	56,88	10,49	93,56	76
AA	Linear Sequence												
163-171 " (mod)	NYATGNLPG NYVTGNIPG	11,16 N/D	3,94 N/D	8,65 N/D	4,69 N/D	1,13 N/D	34,26 N/D	0,51 N/D	9,96 N/D	6,06 N/D	2,33 N/D	6,44 N/D	75 N/D
164-172 " (mod)	YATGNLPGC YVTGNIPGS	8,10 13,33	22,46 6,88	3,67 5,34	2,16 5,18	1,20 2,71	18,82 10,29	0,76 2,70	9,51 12,40	5,09 6,16	2,44 1,98	4,52 4,42	77 88
165-173 " (mod)	ATGNLPGCS VTGNIPGST	7,13 16,18	4,62 8,44	3,64 2,52	2,54 4,85	0,64 1,60	22,84 15,05	1,12 2,24	10,73 13,12	3,81 5,15	2,56 1,58	7,00 4,00	90 105
166-174 " (mod)	TGNLPGCSF TGNIPGSTY	11,97 10,58	5,18 3,59	3,71 13,04	2,56 6,06	4,81 0,86	18,89 35,10	2,38 1,32	8,62 10,25	45,07 52,30	2,37 1,98	11,40 48,40	73 38
167-175 " (mod)	GNLPGCSFS GNIPGSTYS	6,44 7,54	6,11 5,07	3,54 5,04	2,41 1,54	1,13 0,00	13,48 24,38	1,84 0,94	8,71 6,32	3,96 4,80	13,44 1,58	4,80 7,86	80 55
168-176 " (mod)	NLPGCSFSI NIPGSTYSL	13,15 8,41	47,28 5,71	8,15 3,98	3,47 3,20	85,91 8,50	22,34 31,99	1,54 4,80	9,89 7,66	3,11 4,46	3,37 2,29	5,31 4,71	60 37

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171-179 " (mod)	GCSFSIFLL GITFSIYLI	3,65 9,61	5,69 7,27	1,85 1,42	4,77 4,88	2,18 3,05	8,96 10,42	2,88 2,07	11,10 8,94	0,32 0,63	4,28 8,15	2,53 3,29	3,03 9,27	52 68
172-180 " (mod)	CSFSIFLLA ITFSIYLIIV	9,69 9,89	23,38 43,08	5,68 1,66	9,00 5,31	1,37 0,77	6,45 24,10	2,22 4,32	9,85 11,38	0,93 1,53	11,66 6,68	2,99 4,79	4,96 12,30	32 78
173-181 " (mod)	SFSIFLLAL TFSIYLIIVS	5,68 9,93	2,70 2,02	1,42 0,91	2,21 2,89	3,28 0,60	30,35 8,58	1,44 1,28	7,98 9,52	0,60 0,37	8,89 8,63	3,06 1,49	5,33 2,48	21 22
174-182 " (mod)	SRNLGKVID IRNLGRVIE	15,30 14,22	4,94 8,77	3,46 3,89	4,45 5,31	0,71 0,61	9,94 11,06	1,28 1,43	16,71 9,10	0,67 30,77	11,25 4,06	1,28 1,84	4,59 1,98	32 21
175-183 " (mod)	RNLGKVIDT RNLGRVIET	18,90 15,93	6,42 5,96	3,15 5,52	6,22 13,68	1,04 0,51	10,18 15,13	2,42 0,71	13,80 15,51	0,61 0,26	11,79 8,84	2,73 0,91	4,40 2,34	48 27
176-184 " (mod)	NLGKVIDTL NLGRVIETL	18,14 18,24	7,70 11,97	6,41 2,86	12,81 4,53	1,16 0,86	10,67 16,44	2,64 1,47	16,86 15,58	0,57 0,07	12,59 16,29	2,64 1,66	4,39 2,82	52 38
177-185 " (mod)	L GKVIDTLT LGRVIETLT	22,08 16,33	4,05 5,90	7,39 7,40	12,95 20,27	0,70 0,97	15,99 12,31	0,94 2,66	17,96 14,74	0,43 0,28	7,14 16,38	2,90 2,54	5,11 3,43	44 62
178-186 " (mod)	GKVIDTLTC GRVIETLTS	14,77 14,42	4,89 5,26	3,69 12,46	3,96 27,62	0,41 0,88	14,26 15,59	0,25 1,25	13,63 12,91	0,16 0,20	7,31 8,31	0,84 1,05	3,11 3,56	19 38
179-187 " (mod)	KVIDTLTCG RVIELTSG	15,77 11,61	19,13 9,82	4,27 2,23	8,91 3,25	0,57 0,96	13,16 15,52	0,57 0,50	13,17 11,83	0,18 0,11	10,13 37,99	0,63 2,32	2,74 1,90	32 32
180-188	VIDTLTCGF	25,85	5,14	2,93	6,00	0,95	13,90	0,45	10,00	0,11	9,49	1,11	3,26	27

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" (mod)	VIETLTSGF	14,61	4,84	2,55	3,66	0,82	13,07	0,71	12,96	0,35	10,37	1,38	2,39	24
131-139	ADLMGYIPL	31,05	8,06	1,93	3,35	0,61	14,51	0,73	11,40	0,13	8,30	1,01	2,08	26
" (mod)	AEL[Nle]GYIPL	16,63	7,49	3,05	3,61	0,37	20,78	1,11	14,74	0,12	10,20	0,77	1,91	72
132-140	DLMGYIPLV	16,67	103,34	4,66	5,01	1,14	25,64	4,84	22,65	0,23	7,67	1,05	2,49	36
" (mod)	EL[Nle]GYIPLI	18,29	97,87	4,54	3,21	0,92	20,93	0,68	17,05	0,14	8,91	0,67	2,55	42
133-141	LMGYIPLVG	14,17	31,94	2,53	2,13	0,51	24,26	0,41	20,75	0,08	9,43	1,44	0,81	32
" (mod)	L[Nle]GYIPLIG	14,33	49,48	3,98	2,48	0,71	25,94	0,89	17,29	0,15	14,60	0,73	2,85	57
134-142	MGYIPLVGA	11,34	10,71	3,40	4,79	0,35	19,54	0,80	14,88	0,73	8,85	0,83	0,85	40
" (mod)	[Nle]GYIPLIGA	17,87	10,43	3,13	7,27	1,33	27,85	1,78	21,35	0,29	11,07	1,42	5,89	60
135-143	GYIPLVGAP	15,62	7,28	3,46	3,84	0,80	29,74	1,37	15,94	0,47	11,02	1,76	10,35	72
" (mod)	GYLPAVGAP	12,04	5,87	2,90	4,25	0,35	26,03	3,10	17,38	0,11	10,50	0,73	1,84	27
136-144	YIPLVGAPL	21,80	35,00	5,29	3,54	4,51	31,09	41,87	33,10	0,13	54,57	0,69	99,81	30
" (mod)	YLPVAVGAPI	15,76	75,80	2,58	3,07	4,65	28,40	7,70	17,92	0,12	49,63	0,51	2,03	64
137-145	IPLVGAPlG	13,28	4,32	2,59	2,05	0,60	26,79	2,34	17,87	0,03	10,01	0,35	30,74	41
" (mod)	LPAVGAPIG	20,57	5,60	3,02	4,95	0,61	25,32	8,12	15,87	0,12	10,10	0,49	1,06	36
147-155	VARALAHGV	16,10	5,85	2,77	3,47	0,32	29,43	31,59	14,31	0,17	11,26	0,62	0,27	44
" (mod)	VIRVIAHGL	18,66	8,60	4,29	4,54	0,28	27,12	29,93	32,81	0,55	9,64	1,16	2,51	52
148-156	ARALAHGVR	20,62	6,72	2,35	3,83	1,15	31,96	4,70	18,91	1,10	13,58	3,55	2,45	50
" (mod)	IRVIAHGLR	15,82	4,83	3,58	2,50	0,50	23,19	3,50	13,34	0,10	8,99	9,37	0,66	26

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149-157 " (mod)	RALAHGVRV RVIAHGLRL	20,77 11,60	8,69 18,68	2,51 13,32	4,45 17,75	0,84 0,79	29,73 23,22	2,55 102,96	18,08 12,60	0,46 0,42	11,84 106,77	6,37 15,57	6,07 1,43	70 24
27-35 " (mod)	GGQIVGGVY GGQIIIGNY	13,97 24,04	6,88 7,06	3,47 2,23	2,71 2,40	1,78 0,39	20,46 14,38	3,99 1,09	14,66 10,52	0,32 0,25	110,34 90,62	0,66 0,24	0,54 2,22	30 15
28-36 " (mod)	GQIVGGVYL GQIIIGNYL	15,07 20,65	55,85 12,96	4,80 3,01	4,46 2,77	0,50 0,47	13,73 14,17	6,56 2,39	14,53 11,90	0,37 0,22	182,45 173,66	1,01 0,64	3,24 2,60	24 31
29-37 " (mod)	QIVGGVYLL QIIIGNYLI	20,25 21,63	57,04 73,54	2,36 6,30	1,96 4,06	1,01 12,60	13,36 13,60	28,66 52,04	13,15 10,61	0,37 0,40	15,07 29,59	0,60 0,91	4,01 8,99	24 83
30-38 " (mod)	IVGGVYLLP IIGNYLIIP	22,11 21,14	6,78 8,27	1,88 3,88	3,10 2,81	0,30 10,95	12,38 13,88	24,16 15,47	11,48 12,18	1,65 1,17	15,00 19,93	0,68 0,97	2,68 5,35	19 65
31-39 " (mod)	VGGVYLLPR IGGNYLIPR	17,88 17,90	8,73 7,04	3,41 2,88	4,16 3,57	0,34 0,89	14,20 14,85	1,15 20,49	11,57 14,23	0,44 0,32	13,22 15,02	0,79 0,82	4,02 2,54	55 25
41-49 " (mod)	GPRLGVRAT GP[Cit]IGVRAT	13,85 18,60	6,23 9,16	2,45 3,48	1,60 2,72	0,05 0,16	12,66 11,88	124,37 5,38	13,18 9,99	0,94 0,13	10,01 7,29	0,68 0,59	1,83 2,25	19 19
42-50 " (mod)	PRLGVRATR P[Cit]IGVRAT[Cit]	17,77 18,83	6,85 7,36	3,63 2,52	2,73 1,72	0,97 0,21	13,35 14,14	42,64 1,91	11,46 9,47	0,14 0,19	10,04 12,64	0,86 0,94	1,65 2,45	14 45
43-51 " (mod)	RLGVRATRK [Cit]IGVRAT[Cit]R	20,39 19,68	6,71 8,17	112,23 25,16	102,85 8,27	0,28 0,50	12,83 16,67	1,67 14,59	10,21 12,09	0,07 0,17	11,74 14,45	33,13 0,86	2,83 8,51	23 18

Example 4

Intracellular staining:

Peptides according to the invention with X² and X⁴ derived from HCV, Influenza, or CMV were prepared and tested for intracellular staining in an experiment as described above in the “Cell

5 penetration assay”.

Peptide no.	Antigen	X ¹	X ²	X ³	X ⁴ (Different from X ² unless indicated)	X ₅	median	n
1	Negative control		PVHLTLRQAGDDFSR				1.00	
2	Neg control in scaffold	RRG	PVHLTL	RRRG	QAGDDFS		4.12	
3	Positive control		RQIKIWFQNRMMKWKK				2.73	
4	Positive control (tat)		YGRKKRRQRRR				4.43	
5	HCV (Native sequence)		11 amino acids	G	11 amino acids		1.84	
6	HCV	R	11 amino acids of native seq	RRR	11 amino acids of native seq	R	5.35	
7	HCV	R	11 amino acids (1 substitution)	RRR	11 amino acids (5 substitution)	R	11.59	
8	HCV	RR	11 amino acids of native seq	RR	11 amino acids of native seq		6.92	
9	HCV	RR	11 amino acids of native seq	RRR	11 amino acids of native seq		3.25	
10	HCV (Not scaffold)	EE	11 amino acids of native seq	EE	11 amino acids of native seq		1.16	
11	HCV (Not scaffold)	GG	11 amino acids of native seq	GG	11 amino acids of native seq		2.01	
12	HCV (Native sequence)		23 amino acids				3.01	
13	HCV	RR	13 amino acids	RR	10 amino acids of native seq		21.43	
14	HCV	RR	13 amino acids	RR	10 amino acids		4.81	
15	HCV (Not scaffold)	EE	13 amino acids	EE	9 amino acids		1.16	
16	Influenza (Native sequence)		23 amino acids				3.70	
17	Flu	BR	10 amino acids	BRGR	8 amino acids		20.77	
18	cmv	R	9 amino acids	BRR	=X ²	B	3.86	
19	HCV	W	11 amino acids of native seq	RR	11 amino acids of native seq		17.41	
20	HCV	RR	22 amino acids	RR	22 amino acids	R	13.25	

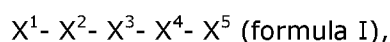
Extracellular staining for the same peptides 1-15:

Peptide no.	Antigen	median	n
1	<i>Negative control</i>	1.00	35
2	Neg control in scaffold	1.77	20
3	Positive control	9.15	19
5	<i>HCV Native sequence</i>	2.08	21
6	HCV	2.20	29
7	HCV	29.42	12
8	HCV	3.06	28
9	HCV	2.06	23
10	<i>HCV (Not scaffold)</i>	1.95	27
11	<i>HCV (Not scaffold)</i>	2.62	23
12	<i>HCV (Native sequence)</i>	1.76	14
13	HCV	16.80	7
14	HCV	17.00	16
15	<i>HCV (Not scaffold)</i>	2.38	12
19	HCV	1.37	19
20	HCV	126.5	7

Results: It was seen that peptides on scaffold form, compared to native peptides or peptides
 5 with other amino acids in X1, X3 and X5 had improved uptake.

Claims

1. An isolated cell-penetrating peptide consisting of the following structure



wherein:

- 5 X^1 and X^3 independently define a linear sequence of any 2, or 3 arginines;
 X^1 and/or X^3 consist of RR or RRR;
 X^2 and X^4 independently define a linear sequence of 8-15 amino acids having at
least 70% sequence identity to an antigen from HCV, CMV, HPV, or Influenza or a
variant sequence thereof;
- 10 the sequence of X^4 is derived from the same antigen as X^2 but is different from
the sequence of X^2 ;
 X^5 is an optional arginine;
wherein the sequence of amino acids defined by $X^1-X^2-X^3-X^4-X^5$ of formula I is not found
in the native sequence of said antigen; and
- 15 wherein the sequence of amino acids defined by $X^1-X^2-X^3-X^4-X^5$ of formula I does not
contain a continuous sequence of four or more arginine residues.
2. The isolated peptide according to claim 1, wherein X^1 and/or X^3 are not part of
the natural sequence of the antigen from which X^2 is derived, and X^3 and/or X^5 are not
part of the natural sequence of the antigen from which X^4 is derived.
- 20 3. The isolated peptide according to either one of claims 1 or 2, wherein the
peptide comprises one or more cysteines.
4. The isolated peptide according to any one of claims 1-3, wherein the N- and/or
C-terminal amino acid in X^2 is a hydrophilic or polar amino acid.
5. The isolated peptide according to any one of claims 1-4, wherein the N-terminal
25 amino acid in X^4 is a hydrophilic or polar amino acid.
6. The isolated peptide according to claim 1, wherein X^1 consists of RR.
7. The isolated peptide according to either one of claims 1 or 6, wherein X^3 consists
of RR.
8. The isolated peptide according to any one of claims 1-7, wherein X^2 and/or X^4 is
30 a linear sequence of less than 12 amino acids.

9. A dimer peptide comprising two peptide monomers, wherein each peptide monomer is as defined in any one of claims 1-8.
10. Peptide combination comprising two or more peptides according to any one of claims 1-9.
- 5 11. An isolated nucleic acid or polynucleotide encoding a peptide according to any one of claims 1-8.
12. A vector comprising the nucleic acid or polynucleotide according to claim 11, or a host cell comprising said vector, with the proviso that said host cell is not a transformed human host cell *in vivo*.
- 10 13. An immunogenic composition comprising at least one peptide according to any one of claims 1-8, a dimer peptide according to claim 9, a peptide combination according to claim 10, the nucleic acid or polynucleotide according to claim 11, or the vector according to claim 12, in combination with a pharmaceutically acceptable diluent or vehicle and optionally an immunological adjuvant.
- 15 14. The immunogenic composition according to claim 13 in the form of a vaccine composition.
15. A peptide according to any one of claims 1-8, a dimer peptide according to claim 9, a peptide combination according to claim 10, the nucleic acid or polynucleotide according to claim 11, the vector according to claim 12, or the immunogenic
- 20 composition according to either one of claims 13 or 14 for use in a method of inducing an immune response in a subject against an antigen.
16. A peptide according to any one of claims 1-8, a dimer peptide according to claim 9, a peptide combination according to claim 10, the nucleic acid or polynucleotide according to claim 11, the vector according to claim 12, or the immunogenic
- 25 composition according to either one of claims 13 or 14 for use in a method of reducing and/or delaying the pathological effects of a virus in a subject infected with said virus.
17. A peptide according to any one of claims 1-9 for use as a medicament.
18. A peptide according to any one of claims 1-9 when used the pathological effects of a virus in a subject infected with said virus.

19. A method of producing the isolated cell penetrating peptide of any one of claims 1-8, the method comprising:

providing a first antigen X^2 and a second antigen X^4 , wherein X^2 and X^4 independently define a linear sequence of 8-15 amino acids having at least 70% sequence identity to an antigen from HCV, CMV, HPV, or Influenza or a variant sequence thereof, and wherein the sequence of X^4 is derived from the same antigen as X^2 but is different from the sequence of X^2 ; and

inserting a sequence of arginines at each end of X^2 and X^4 such that the resulting peptide conforms to the following structure

X^1 - X^2 - X^3 - X^4 - X^5 (formula I),

wherein:

X^1 and X^3 independently define a linear sequence of any 2, or 3 arginines;

X^1 and/or X^3 consist of RR or RRR;

X^5 is an optional arginine;

wherein the sequence of amino acids defined by X^1 - X^2 - X^3 - X^4 - X^5 of formula I is not found in the native sequence of said antigen; and

wherein the sequence of amino acids defined by X^1 - X^2 - X^3 - X^4 - X^5 of formula I does not contain a continuous sequence of four or more arginine residues.

20. The method of claim 21, wherein X^1 and/or X^3 are not part of the natural

sequence of the antigen from which X^2 is derived, and X^3 and/or X^5 are not part of the natural sequence of the antigen from which X^4 is derived.

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Page 1

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Val	Pro	Gln	Ser	Phe	Gln	Val	Ala	His	Leu	His	Ala	Pro	Thr	Gly	
1220						1225					1230				
Ser	Gly	Lys	Ser	Thr	Lys	Val	Pro	Ala	Ala	Tyr	Ala	Ala	Gln	Gly	
1235						1240					1245				
Tyr	Lys	Val	Leu	Val	Leu	Asn	Pro	Ser	Val	Ala	Ala	Thr	Leu	Gly	
1250						1255					1260				
Phe	Gly	Ala	Tyr	Met	Ser	Lys	Ala	His	Gly	Val	Asp	Pro	Asn	Ile	

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1265						1270								
Arg	Thr	Gly	Val	Arg	Thr	Ile	Thr	Thr	Gly	Ser	Pro	Ile	Thr	Tyr
1280						1285					1290			
Ser	Thr	Tyr	Gly	Lys	Phe	Leu	Ala	Asp	Gly	Gly	Cys	Ser	Gly	Gly
1295						1300					1305			
Ala	Tyr	Asp	Ile	Ile	Ile	Cys	Asp	Glu	Cys	His	Ser	Thr	Asp	Ala
1310						1315					1320			
Thr	Ser	Ile	Leu	Gly	Ile	Gly	Thr	Val	Leu	Asp	Gln	Ala	Glu	Thr
1325						1330					1335			
Ala	Gly	Ala	Arg	Leu	Val	Val	Leu	Ala	Thr	Ala	Thr	Pro	Pro	Gly
1340						1345					1350			
Ser	Val	Thr	Val	Ser	His	Pro	Asn	Ile	Glu	Glu	Val	Ala	Leu	Ser
1355						1360					1365			
Thr	Thr	Gly	Glu	Ile	Pro	Phe	Tyr	Gly	Lys	Ala	Ile	Pro	Leu	Glu
1370						1375					1380			
Val	Ile	Lys	Gly	Gly	Arg	His	Leu	Ile	Phe	Cys	His	Ser	Lys	Lys
1385						1390					1395			
Lys	Cys	Asp	Glu	Leu	Ala	Ala	Lys	Leu	Val	Ala	Leu	Gly	Ile	Asn
1400						1405					1410			
Ala	Val	Ala	Tyr	Tyr	Arg	Gly	Leu	Asp	Val	Ser	Val	Ile	Pro	Thr
1415						1420					1425			
Ser	Gly	Asp	Val	Val	Val	Val	Ser	Thr	Asp	Ala	Leu	Met	Thr	Gly
1430						1435					1440			
Phe	Thr	Gly	Asp	Phe	Asp	Ser	Val	Ile	Asp	Cys	Asn	Thr	Cys	Val
1445						1450					1455			
Thr	Gln	Thr	Val	Asp	Phe	Ser	Leu	Asp	Pro	Thr	Phe	Thr	Ile	Glu
1460						1465					1470			
Thr	Thr	Thr	Leu	Pro	Gln	Asp	Ala	Val	Ser	Arg	Thr	Gln	Arg	Arg
1475						1480					1485			
Gly	Arg	Thr	Gly	Arg	Gly	Lys	Pro	Gly	Ile	Tyr	Arg	Phe	Val	Ala
1490						1495					1500			
Pro	Gly	Glu	Arg	Pro	Ser	Gly	Met	Phe	Asp	Ser	Ser	Val	Leu	Cys
1505						1510					1515			
Glu	Cys	Tyr	Asp	Ala	Gly	Cys	Ala	Trp	Tyr	Glu	Leu	Thr	Pro	Ala

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1520						1525						1530			
Glu	Thr	Thr	Val	Arg	Leu	Arg	Ala	Tyr	Met	Asn	Thr	Pro	Gly	Leu	
	1535					1540					1545				
Pro	Val	Cys	Gln	Asp	His	Leu	Glu	Phe	Trp	Glu	Gly	Val	Phe	Thr	
	1550					1555					1560				
Gly	Leu	Thr	His	Ile	Asp	Ala	His	Phe	Leu	Ser	Gln	Thr	Lys	Gln	
	1565					1570					1575				
Ser	Gly	Glu	Asn	Phe	Pro	Tyr	Leu	Val	Ala	Tyr	Gln	Ala	Thr	Val	
	1580					1585					1590				
Cys	Ala	Arg	Ala	Gln	Ala	Pro	Pro	Pro	Ser	Trp	Asp	Gln	Met	Trp	
	1595					1600					1605				
Lys	Cys	Leu	Ile	Arg	Leu	Lys	Pro	Thr	Leu	His	Gly	Pro	Thr	Pro	
	1610					1615					1620				
Leu	Leu	Tyr	Arg	Leu	Gly	Ala	Val	Gln	Asn	Glu	Val	Thr	Leu	Thr	
	1625					1630					1635				
His	Pro	Ile	Thr	Lys	Tyr	Ile	Met	Thr	Cys	Met	Ser	Ala	Asp	Leu	
	1640					1645					1650				
Glu	Val	Val	Thr	Ser	Thr	Trp	Val	Leu	Val	Gly	Gly	Val	Leu	Ala	
	1655					1660					1665				
Ala	Leu	Ala	Ala	Tyr	Cys	Leu	Ser	Thr	Gly	Cys	Val	Val	Ile	Val	
	1670					1675					1680				
Gly	Arg	Ile	Val	Leu	Ser	Gly	Lys	Pro	Ala	Ile	Ile	Pro	Asp	Arg	
	1685					1690					1695				
Glu	Val	Leu	Tyr	Gln	Glu	Phe	Asp	Glu	Met	Glu	Glu	Cys	Ser	Gln	
	1700					1705					1710				
His	Leu	Pro	Tyr	Ile	Glu	Gln	Gly	Met	Met	Leu	Ala	Glu	Gln	Phe	
	1715					1720					1725				
Lys	Gln	Lys	Ala	Leu	Gly	Leu	Leu	Gln	Thr	Ala	Ser	Arg	Gln	Ala	
	1730					1735					1740				
Glu	Val	Ile	Thr	Pro	Ala	Val	Gln	Thr	Asn	Trp	Gln	Lys	Leu	Glu	
	1745					1750					1755				
Val	Phe	Trp	Ala	Lys	His	Met	Trp	Asn	Phe	Ile	Ser	Gly	Ile	Gln	
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Tyr	Leu	Ala	Gly	Leu	Ser	Thr	Leu	Pro	Gly	Asn	Pro	Ala	Ile	Ala	

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1775						1780								
Ser	Leu	Met	Ala	Phe	Thr	Ala	Ala	Val	Thr	Ser	Pro	Leu	Thr	Thr
1790						1795					1800			
Gly	Gln	Thr	Leu	Leu	Phe	Asn	Ile	Leu	Gly	Gly	Trp	Val	Ala	Ala
1805						1810					1815			
Gln	Leu	Ala	Ala	Pro	Gly	Ala	Ala	Thr	Ala	Phe	Val	Gly	Ala	Gly
1820						1825					1830			
Leu	Ala	Gly	Ala	Ala	Ile	Gly	Ser	Val	Gly	Leu	Gly	Lys	Val	Leu
1835						1840					1845			
Val	Asp	Ile	Leu	Ala	Gly	Tyr	Gly	Ala	Gly	Val	Ala	Gly	Ala	Leu
1850						1855					1860			
Val	Ala	Phe	Lys	Ile	Met	Ser	Gly	Glu	Val	Pro	Ser	Thr	Glu	Asp
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Leu	Val	Asn	Leu	Leu	Pro	Ala	Ile	Leu	Ser	Pro	Gly	Ala	Leu	Val
1880						1885					1890			
Val	Gly	Val	Val	Cys	Ala	Ala	Ile	Leu	Arg	Arg	His	Val	Gly	Pro
1895						1900					1905			
Gly	Glu	Gly	Ala	Val	Gln	Trp	Met	Asn	Arg	Leu	Ile	Ala	Phe	Ala
1910						1915					1920			
Ser	Arg	Gly	Asn	His	Val	Ser	Pro	Thr	His	Tyr	Val	Pro	Glu	Ser
1925						1930					1935			
Asp	Ala	Ala	Ala	Arg	Val	Thr	Ala	Ile	Leu	Ser	Ser	Leu	Thr	Val
1940						1945					1950			
Thr	Gln	Leu	Leu	Arg	Arg	Leu	His	Gln	Trp	Ile	Ser	Ser	Glu	Cys
1955						1960					1965			
Thr	Thr	Pro	Cys	Ser	Gly	Ser	Trp	Leu	Arg	Asp	Ile	Trp	Asp	Trp
1970						1975					1980			
Ile	Cys	Glu	Val	Leu	Ser	Asp	Phe	Lys	Thr	Trp	Leu	Lys	Ala	Lys
1985						1990					1995			
Leu	Met	Pro	Gln	Leu	Pro	Gly	Ile	Pro	Phe	Val	Ser	Cys	Gln	Arg
2000						2005					2010			
Gly	Tyr	Arg	Gly	Val	Trp	Arg	Gly	Asp	Gly	Ile	Met	His	Thr	Arg
2015						2020					2025			
Cys	His	Cys	Gly	Ala	Glu	Ile	Thr	Gly	His	Val	Lys	Asn	Gly	Thr

2030						2035						2040			
Met	Arg	Ile	Val	Gly	Pro	Arg	Thr	Cys	Arg	Asn	Met	Trp	Ser	Gly	
	2045					2050					2055				
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	2060					2065					2070				
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	2090					2095					2100				
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	2105					2110					2115				
Pro	Glu	Phe	Phe	Thr	Glu	Leu	Asp	Gly	Val	Arg	Leu	His	Arg	Phe	
	2120					2125					2130				
Ala	Pro	Pro	Cys	Lys	Pro	Leu	Leu	Arg	Glu	Glu	Val	Ser	Phe	Arg	
	2135					2140					2145				
Val	Gly	Leu	His	Glu	Tyr	Pro	Val	Gly	Ser	Gln	Leu	Pro	Cys	Glu	
	2150					2155					2160				
Pro	Glu	Pro	Asp	Val	Ala	Val	Leu	Thr	Ser	Met	Leu	Thr	Asp	Pro	
	2165					2170					2175				
Ser	His	Ile	Thr	Ala	Glu	Ala	Ala	Gly	Arg	Arg	Leu	Ala	Arg	Gly	
	2180					2185					2190				
Ser	Pro	Pro	Ser	Met	Ala	Ser	Ser	Ser	Ala	Ser	Gln	Leu	Ser	Ala	
	2195					2200					2205				
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Ala	Glu	Leu	Ile	Glu	Ala	Asn	Leu	Leu	Trp	Arg	Gln	Glu	Met	Gly	
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Gly	Asn	Ile	Thr	Arg	Val	Glu	Ser	Glu	Asn	Lys	Val	Val	Ile	Leu	
	2240					2245					2250				
Asp	Ser	Phe	Asp	Pro	Leu	Val	Ala	Glu	Glu	Asp	Glu	Arg	Glu	Val	
	2255					2260					2265				
Ser	Val	Pro	Ala	Glu	Ile	Leu	Arg	Lys	Ser	Arg	Arg	Phe	Ala	Arg	
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Ala	Leu	Pro	Val	Trp	Ala	Arg	Pro	Asp	Tyr	Asn	Pro	Pro	Leu	Val	

2295

Ile Asn Ser Val Trp Lys Asp Leu Leu Glu Asp Ser Val Thr Pro

2540						2545								
Ile	Asp	Thr	Thr	Ile	Met	Ala	Lys	Asn	Glu	Val	Phe	Cys	Val	Gln
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Pro	Glu	Lys	Gly	Gly	Arg	Lys	Pro	Ala	Arg	Leu	Ile	Val	Phe	Pro
	2570					2575					2580			
Asp	Leu	Gly	Val	Arg	Val	Cys	Glu	Lys	Met	Ala	Leu	Tyr	Asp	Val
	2585					2590					2595			
Val	Ser	Lys	Leu	Pro	Leu	Ala	Val	Met	Gly	Ser	Ser	Tyr	Gly	Phe
	2600					2605					2610			
Gln	Tyr	Ser	Pro	Gly	Gln	Arg	Val	Glu	Phe	Leu	Val	Gln	Ala	Trp
	2615					2620					2625			
Lys	Ser	Lys	Lys	Thr	Pro	Met	Gly	Phe	Ser	Tyr	Asp	Thr	Arg	Cys
	2630					2635					2640			
Phe	Asp	Ser	Thr	Val	Thr	Glu	Ser	Asp	Ile	Arg	Thr	Glu	Glu	Ala
	2645					2650					2655			
Ile	Tyr	Gln	Cys	Cys	Asp	Leu	Asp	Pro	Gln	Ala	Arg	Val	Ala	Ile
	2660					2665					2670			
Lys	Ser	Leu	Thr	Glu	Arg	Leu	Tyr	Val	Gly	Gly	Pro	Leu	Thr	Asn
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Ser	Arg	Gly	Glu	Asn	Cys	Gly	Tyr	Arg	Arg	Cys	Arg	Ala	Ser	Gly
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Ala	Arg	Ala	Ala	Cys	Arg	Ala	Ala	Gly	Leu	Gln	Asp	Cys	Thr	Met
	2720					2725					2730			
Leu	Val	Cys	Gly	Asp	Asp	Leu	Val	Val	Ile	Cys	Glu	Ser	Ala	Gly
	2735					2740					2745			
Val	Gln	Glu	Asp	Ala	Ala	Ser	Leu	Arg	Ala	Phe	Thr	Glu	Ala	Met
	2750					2755					2760			
Thr	Arg	Tyr	Ser	Ala	Pro	Pro	Gly	Asp	Pro	Pro	Gln	Pro	Glu	Tyr
	2765					2770					2775			
Asp	Leu	Glu	Leu	Ile	Thr	Ser	Cys	Ser	Ser	Asn	Val	Ser	Val	Ala
	2780					2785					2790			
His	Asp	Gly	Ala	Gly	Lys	Arg	Val	Tyr	Tyr	Leu	Thr	Arg	Asp	Pro

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2795

2800

2805

Thr	Thr	Pro	Leu	Ala	Arg	Ala	Ala	Trp	Glu	Thr	Ala	Arg	His	Thr
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Pro	Val	Asn	Ser	Trp	Leu	Gly	Asn	Ile	Ile	Met	Phe	Ala	Pro	Thr
	2825					2830					2835			
Leu	Trp	Ala	Arg	Met	Ile	Leu	Met	Thr	His	Phe	Phe	Ser	Val	Leu
	2840					2845					2850			
Ile	Ala	Arg	Asp	Gln	Leu	Glu	Gln	Ala	Leu	Asn	Cys	Glu	Ile	Tyr
	2855					2860					2865			
Gly	Ala	Cys	Tyr	Ser	Ile	Glu	Pro	Leu	Asp	Leu	Pro	Pro	Ile	Ile
	2870					2875					2880			
Gln	Arg	Leu	His	Gly	Leu	Ser	Ala	Phe	Ser	Leu	His	Ser	Tyr	Ser
	2885					2890					2895			
Pro	Gly	Glu	Ile	Asn	Arg	Val	Ala	Ala	Cys	Leu	Arg	Lys	Leu	Gly
	2900					2905					2910			
Val	Pro	Pro	Leu	Arg	Ala	Trp	Arg	His	Arg	Ala	Arg	Ser	Val	Arg
	2915					2920					2925			
Ala	Arg	Leu	Leu	Ser	Arg	Gly	Gly	Arg	Ala	Ala	Ile	Cys	Gly	Lys
	2930					2935					2940			
Tyr	Leu	Phe	Asn	Trp	Ala	Val	Arg	Thr	Lys	Leu	Lys	Leu	Thr	Pro
	2945					2950					2955			
Ile	Ala	Ala	Ala	Gly	Arg	Leu	Asp	Leu	Ser	Gly	Trp	Phe	Thr	Ala
	2960					2965					2970			
Gly	Tyr	Ser	Gly	Gly	Asp	Ile	Tyr	His	Ser	Val	Ser	His	Ala	Arg
	2975					2980					2985			
Pro	Arg	Trp	Phe	Trp	Phe	Cys	Leu	Leu	Leu	Leu	Ala	Ala	Gly	Val
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Gly	Ile	Tyr	Leu	Leu	Pro	Asn	Arg							
	3005					3010								

<210> 2

<211> 191

<212> PRT

<213> Hepatitis C virus

<400> 2

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Arg Arg Pro Gln Asp Val Lys Phe Pro Gly Gly Gly Gln Ile Val Gly
20 25 30

Gly Val Tyr Leu Leu Pro Arg Arg Gly Pro Arg Leu Gly Val Arg Ala
35 40 45

Thr Arg Lys Thr Ser Glu Arg Ser Gln Pro Arg Gly Arg Arg Gln Pro
50 55 60

Ile Pro Lys Ala Arg Arg Pro Glu Gly Arg Thr Trp Ala Gln Pro Gly
65 70 75 80

Tyr Pro Trp Pro Leu Tyr Gly Asn Glu Gly Cys Gly Trp Ala Gly Trp
85 90 95

Leu Leu Ser Pro Arg Gly Ser Arg Pro Ser Trp Gly Pro Thr Asp Pro
100 105 110

Arg Arg Arg Ser Arg Asn Leu Gly Lys Val Ile Asp Thr Leu Thr Cys
115 120 125

Gly Phe Ala Asp Leu Met Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu
130 135 140

Gly Gly Ala Ala Arg Ala Leu Ala His Gly Val Arg Val Leu Glu Asp
145 150 155 160

Gly Val Asn Tyr Ala Thr Gly Asn Leu Pro Gly Cys Ser Phe Ser Ile
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Phe Leu Leu Ala Leu Leu Ser Cys Leu Thr Val Pro Ala Ser Ala
180 185 190

<210> 3
<211> 3010
<212> PRT
<213> Hepatitis C virus

<400> 3

Met Ser Thr Asn Pro Lys Pro Gln Arg Lys Thr Lys Arg Asn Thr Asn
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20 25 30

Gly Val Tyr Leu Leu Pro Arg Arg Gly Pro Arg Leu Gly Val Arg Ala
35 40 45

Thr Arg Lys Thr Ser Glu Arg Ser Gln Pro Arg Gly Arg Arg Gln Pro
50 55 60

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Ile Pro Lys Ala Arg Arg Pro Glu Gly Arg Ala Trp Ala Gln Pro Gly
65      70      75      80

Tyr Pro Trp Pro Leu Tyr Gly Asn Glu Gly Leu Gly Trp Ala Gly Trp
      85      90      95

Leu Leu Ser Pro Arg Gly Ser Arg Pro Ser Trp Gly Pro Thr Asp Pro
      100      105      110

Arg Arg Lys Ser Arg Asn Leu Gly Lys Val Ile Asp Thr Leu Thr Cys
      115      120      125

Gly Phe Ala Asp Leu Met Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu
      130      135      140

Gly Gly Val Ala Arg Ala Leu Ala His Gly Val Arg Val Leu Glu Asp
145      150      155      160

Gly Val Asn Tyr Ala Thr Gly Asn Leu Pro Gly Cys Ser Phe Ser Ile
      165      170      175

Phe Leu Leu Ala Leu Leu Ser Cys Leu Thr Thr Pro Val Ser Ala Tyr
      180      185      190

Glu Val Arg Asn Ala Ser Gly Met Tyr His Val Thr Asn Asp Cys Ser
      195      200      205

Asn Ser Ser Ile Val Tyr Glu Ala Ala Asp Met Ile Met His Thr Pro
      210      215      220

Gly Cys Val Pro Cys Val Arg Glu Asp Asn Ser Ser Arg Cys Trp Val
225      230      235      240

Ala Leu Thr Pro Thr Leu Ala Ala Arg Asn Ala Ser Val Pro Thr Thr
      245      250      255

Thr Leu Arg Arg His Val Asp Leu Leu Val Gly Val Ala Ala Phe Cys
      260      265      270

Ser Ala Met Tyr Val Gly Asp Leu Cys Gly Ser Val Phe Leu Val Ser
      275      280      285

Gln Leu Phe Thr Phe Ser Pro Arg Arg His Glu Thr Val Gln Asp Cys
      290      295      300

Asn Cys Ser Ile Tyr Pro Gly Arg Val Ser Gly His Arg Met Ala Trp
305      310      315      320

Asp Met Met Met Asn Trp Ser Pro Thr Thr Ala Leu Val Val Ser Gln
      325      330      335

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Leu Leu Arg Ile Pro Gln Ala Val Val Asp Met Val Thr Gly Ser His
 340 345 350
 Trp Gly Ile Leu Ala Gly Leu Ala Tyr Tyr Ser Met Val Gly Asn Trp
 355 360 365
 Ala Lys Val Leu Ile Ala Met Leu Leu Phe Ala Gly Val Asp Gly Thr
 370 375 380
 Thr His Val Thr Gly Gly Ala Gln Gly Arg Ala Ala Ser Ser Leu Thr
 385 390 395 400
 Ser Leu Phe Ser Pro Gly Pro Val Gln His Leu Gln Leu Ile Asn Thr
 405 410 415
 Asn Gly Ser Trp His Ile Asn Arg Thr Ala Leu Ser Cys Asn Asp Ser
 420 425 430
 Leu Asn Thr Gly Phe Val Ala Ala Leu Phe Tyr Lys Tyr Arg Phe Asn
 435 440 445
 Ala Ser Gly Cys Pro Glu Arg Leu Ala Thr Cys Arg Pro Ile Asp Thr
 450 455 460
 Phe Ala Gln Gly Trp Gly Pro Ile Thr Tyr Thr Glu Pro His Asp Leu
 465 470 475 480
 Asp Gln Arg Pro Tyr Cys Trp His Tyr Ala Pro Gln Pro Cys Gly Ile
 485 490 495
 Val Pro Thr Leu Gln Val Cys Gly Pro Val Tyr Cys Phe Thr Pro Ser
 500 505 510
 Pro Val Ala Val Gly Thr Thr Asp Arg Phe Gly Ala Pro Thr Tyr Arg
 515 520 525
 Trp Gly Ala Asn Glu Thr Asp Val Leu Leu Leu Asn Asn Ala Gly Pro
 530 535 540
 Pro Gln Gly Asn Trp Phe Gly Cys Thr Trp Met Asn Gly Thr Gly Phe
 545 550 555 560
 Thr Lys Thr Cys Gly Gly Pro Pro Cys Asn Ile Gly Gly Val Gly Asn
 565 570 575
 Asn Thr Leu Thr Cys Pro Thr Asp Cys Phe Arg Lys His Pro Gly Ala
 580 585 590
 Thr Tyr Thr Lys Cys Gly Ser Gly Pro Trp Leu Thr Pro Arg Cys Leu
 595 600 605

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Val	Asp	Tyr	Pro	Tyr	Arg	Leu	Trp	His	Tyr	Pro	Cys	Thr	Val	Asn	Phe		
	610					615					620						
Thr	Ile	Phe	Lys	Val	Arg	Met	Tyr	Val	Gly	Gly	Ala	Glu	His	Arg	Leu		
625					630					635					640		
Asp	Ala	Ala	Cys	Asn	Trp	Thr	Arg	Gly	Glu	Arg	Cys	Asp	Leu	Glu	Asp		
				645					650					655			
Arg	Asp	Arg	Ser	Glu	Leu	Ser	Pro	Leu	Leu	Leu	Ser	Thr	Thr	Glu	Trp		
			660					665					670				
Gln	Val	Leu	Pro	Cys	Ser	Phe	Thr	Thr	Leu	Pro	Ala	Leu	Ser	Thr	Gly		
		675					680					685					
Leu	Ile	His	Leu	His	Gln	Asn	Ile	Val	Asp	Ile	Gln	Tyr	Leu	Tyr	Gly		
	690					695					700						
Ile	Gly	Ser	Ala	Val	Val	Ser	Phe	Ala	Ile	Lys	Trp	Glu	Tyr	Ile	Val		
705					710					715					720		
Leu	Leu	Phe	Leu	Leu	Leu	Ala	Asp	Ala	Arg	Val	Cys	Ala	Cys	Leu	Trp		
				725					730					735			
Met	Met	Leu	Leu	Val	Ala	Gln	Ala	Glu	Ala	Ala	Leu	Glu	Asn	Leu	Val		
			740					745					750				
Val	Leu	Asn	Ala	Ala	Ser	Val	Ala	Gly	Ala	His	Gly	Ile	Leu	Ser	Phe		
		755					760					765					
Ile	Val	Phe	Phe	Cys	Ala	Ala	Trp	Tyr	Ile	Lys	Gly	Arg	Leu	Val	Pro		
	770					775					780						
Gly	Ala	Ala	Tyr	Ala	Leu	Tyr	Gly	Val	Trp	Pro	Leu	Leu	Leu	Leu	Leu		
785					790					795					800		
Leu	Ala	Leu	Pro	Pro	Arg	Ala	Tyr	Ala	Met	Asp	Arg	Glu	Met	Ala	Ala		
				805					810					815			
Ser	Cys	Gly	Gly	Ala	Val	Phe	Val	Gly	Leu	Val	Leu	Leu	Thr	Leu	Ser		
			820					825					830				
Pro	His	Tyr	Lys	Val	Phe	Leu	Ala	Arg	Phe	Ile	Trp	Trp	Leu	Gln	Tyr		
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Leu	Ile	Thr	Arg	Thr	Glu	Ala	His	Leu	Gln	Val	Trp	Val	Pro	Pro	Leu		
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Asn	Val	Arg	Gly	Gly	Arg	Asp	Ala	Ile	Ile	Leu	Leu	Thr	Cys	Val	Val		
865					870					875					880		

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His Pro Glu Leu Ile Phe Asp Ile Thr Lys Tyr Leu Leu Ala Ile Phe
885 890 895
Gly Pro Leu Met Val Leu Gln Ala Gly Ile Thr Arg Val Pro Tyr Phe
900 905 910
Val Arg Ala Gln Gly Leu Ile Arg Ala Cys Met Leu Ala Arg Lys Val
915 920 925
Val Gly Gly His Tyr Val Gln Met Val Phe Met Lys Leu Ala Ala Leu
930 935 940
Ala Gly Thr Tyr Val Tyr Asp His Leu Thr Pro Leu Arg Asp Trp Ala
945 950 955 960
His Thr Gly Leu Arg Asp Leu Ala Val Ala Val Glu Pro Val Val Phe
965 970 975
Ser Asp Met Glu Thr Lys Val Ile Thr Trp Gly Ala Asp Thr Ala Ala
980 985 990
Cys Gly Asp Ile Ile Leu Ala Leu Pro Ala Ser Ala Arg Arg Gly Lys
995 1000 1005
Glu Ile Leu Leu Gly Pro Ala Asp Ser Leu Glu Gly Gln Gly Trp
1010 1015 1020
Arg Leu Leu Ala Pro Ile Thr Ala Tyr Ser Gln Gln Thr Arg Gly
1025 1030 1035
Leu Leu Gly Cys Ile Ile Thr Ser Leu Thr Gly Arg Asp Lys Asn
1040 1045 1050
Gln Val Glu Gly Glu Val Gln Val Val Ser Thr Ala Thr Gln Ser
1055 1060 1065
Phe Leu Ala Thr Cys Ile Asn Gly Val Cys Trp Thr Val Phe His
1070 1075 1080
Gly Ala Gly Ser Lys Thr Leu Ala Gly Pro Lys Gly Pro Ile Thr
1085 1090 1095
Gln Met Tyr Thr Asn Val Asp Gln Asp Leu Val Gly Trp Pro Ala
1100 1105 1110
Pro Pro Gly Ala Arg Ser Leu Thr Pro Cys Thr Cys Gly Ser Ser
1115 1120 1125
Asp Leu Tyr Leu Val Thr Arg His Ala Asp Val Ile Pro Val Arg
1130 1135 1140

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Arg	Arg 1145	Gly	Asp	Gly	Arg	Gly 1150	Ser	Leu	Leu	Pro	Pro 1155	Arg	Pro	Val
Ser	Tyr 1160	Leu	Lys	Gly	Ser	Ser 1165	Gly	Gly	Pro	Leu	Leu 1170	Cys	Pro	Ser
Gly	His 1175	Ala	Val	Gly	Ile	Leu 1180	Pro	Ala	Ala	Val	Cys 1185	Thr	Arg	Gly
Val	Ala 1190	Met	Ala	Val	Glu	Phe 1195	Ile	Pro	Val	Glu	Ser 1200	Met	Glu	Thr
Thr	Met 1205	Arg	Ser	Pro	Val	Phe 1210	Thr	Asp	Asn	Pro	Ser 1215	Pro	Pro	Ala
Val	Pro 1220	Gln	Thr	Phe	Gln	Val 1225	Ala	His	Leu	His	Ala 1230	Pro	Thr	Gly
Ser	Gly 1235	Lys	Ser	Thr	Arg	Val 1240	Pro	Ala	Ala	Tyr	Ala 1245	Ala	Gln	Gly
Tyr	Lys 1250	Val	Leu	Val	Leu	Asn 1255	Pro	Ser	Val	Ala	Ala 1260	Thr	Leu	Gly
Phe	Gly 1265	Ala	Tyr	Met	Ser	Lys 1270	Ala	His	Gly	Ile	Asp 1275	Pro	Asn	Leu
Arg	Thr 1280	Gly	Val	Arg	Thr	Ile 1285	Thr	Thr	Gly	Ala	Pro 1290	Ile	Thr	Tyr
Ser	Thr 1295	Tyr	Gly	Lys	Phe	Leu 1300	Ala	Asp	Gly	Gly	Gly 1305	Ser	Gly	Gly
Ala	Tyr 1310	Asp	Ile	Ile	Met	Cys 1315	Asp	Glu	Cys	His	Ser 1320	Thr	Asp	Ser
Thr	Thr 1325	Ile	Tyr	Gly	Ile	Gly 1330	Thr	Val	Leu	Asp	Gln 1335	Ala	Glu	Thr
Ala	Gly 1340	Ala	Arg	Leu	Val	Val 1345	Leu	Ser	Thr	Ala	Thr 1350	Pro	Pro	Gly
Ser	Val 1355	Thr	Val	Pro	His	Leu 1360	Asn	Ile	Glu	Glu	Val 1365	Ala	Leu	Ser
Asn	Thr 1370	Gly	Glu	Ile	Pro	Phe 1375	Tyr	Gly	Lys	Ala	Ile 1380	Pro	Ile	Glu
Ala	Ile 1385	Lys	Gly	Gly	Arg	His 1390	Leu	Ile	Phe	Cys	His 1395	Ser	Lys	Lys

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Lys	Cys 1400	Asp	Glu	Leu	Ala	Ala 1405	Lys	Leu	Ser	Gly	Leu 1410	Gly	Leu	Asn
Ala	Val 1415	Ala	Tyr	Tyr	Arg	Gly 1420	Leu	Asp	Val	Ser	Val 1425	Ile	Pro	Thr
Ser	Gly 1430	Asp	Val	Val	Val	Val 1435	Ala	Thr	Asp	Ala	Leu 1440	Met	Thr	Gly
Phe	Thr 1445	Gly	Asp	Phe	Asp	Ser 1450	Val	Ile	Asp	Cys	Asn 1455	Thr	Cys	Val
Thr	Gln 1460	Thr	Val	Asp	Phe	Ser 1465	Leu	Asp	Pro	Thr	Phe 1470	Thr	Ile	Glu
Thr	Thr 1475	Thr	Val	Pro	Gln	Asp 1480	Ala	Val	Ser	Arg	Ser 1485	Gln	Arg	Arg
Gly	Arg 1490	Thr	Gly	Arg	Gly	Arg 1495	Ala	Gly	Ile	Tyr	Arg 1500	Phe	Val	Thr
Pro	Gly 1505	Glu	Arg	Pro	Ser	Gly 1510	Met	Phe	Asp	Ser	Ser 1515	Val	Leu	Cys
Glu	Cys 1520	Tyr	Asp	Ala	Gly	Cys 1525	Ala	Trp	Tyr	Glu	Leu 1530	Thr	Pro	Ala
Glu	Thr 1535	Ser	Val	Arg	Leu	Arg 1540	Ala	Tyr	Leu	Asn	Thr 1545	Pro	Gly	Leu
Pro	Val 1550	Cys	Gln	Asp	His	Leu 1555	Glu	Phe	Ser	Glu	Gly 1560	Val	Phe	Thr
Gly	Leu 1565	Thr	His	Ile	Asp	Ala 1570	His	Phe	Leu	Ser	Gln 1575	Thr	Lys	Gln
Ala	Gly 1580	Glu	Asn	Phe	Pro	Tyr 1585	Leu	Val	Ala	Tyr	Gln 1590	Ala	Thr	Val
Cys	Ala 1595	Arg	Ala	Gln	Ala	Pro 1600	Pro	Pro	Ser	Trp	Asp 1605	Glu	Met	Trp
Arg	Cys 1610	Leu	Ile	Arg	Leu	Lys 1615	Pro	Thr	Leu	His	Gly 1620	Pro	Thr	Pro
Leu	Leu 1625	Tyr	Arg	Leu	Gly	Ala 1630	Val	Gln	Asn	Glu	Val 1635	Thr	Leu	Thr
His	Pro 1640	Ile	Thr	Lys	Phe	Ile 1645	Met	Thr	Cys	Met	Ser 1650	Ala	Asp	Leu

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Glu	Val	Val	Thr	Ser	Thr	Trp	Val	Leu	Val	Gly	Gly	Val	Leu	Ala
1655						1660					1665			
Ala	Leu	Ala	Ala	Tyr	Cys	Leu	Thr	Thr	Gly	Ser	Val	Val	Ile	Val
1670						1675					1680			
Gly	Arg	Ile	Ile	Leu	Ser	Gly	Lys	Pro	Ala	Ile	Ile	Pro	Asp	Arg
1685						1690					1695			
Glu	Val	Leu	Tyr	Gln	Glu	Phe	Asp	Glu	Met	Glu	Glu	Cys	Ala	Ser
1700						1705					1710			
His	Leu	Pro	Tyr	Phe	Glu	Gln	Gly	Met	Gln	Leu	Ala	Glu	Gln	Phe
1715						1720					1725			
Lys	Gln	Lys	Ala	Leu	Gly	Leu	Leu	Gln	Thr	Ala	Thr	Lys	Gln	Ala
1730						1735					1740			
Glu	Ala	Ala	Ala	Pro	Val	Val	Glu	Ser	Lys	Trp	Arg	Ala	Leu	Glu
1745						1750					1755			
Thr	Phe	Trp	Ala	Lys	His	Met	Trp	Asn	Phe	Ile	Ser	Gly	Ile	Gln
1760						1765					1770			
Tyr	Leu	Ala	Gly	Leu	Ser	Thr	Leu	Pro	Gly	Asn	Pro	Ala	Ile	Arg
1775						1780					1785			
Ser	Pro	Met	Ala	Phe	Thr	Ala	Ser	Ile	Thr	Ser	Pro	Leu	Thr	Thr
1790						1795					1800			
Gln	His	Thr	Leu	Leu	Phe	Asn	Ile	Leu	Gly	Gly	Trp	Val	Ala	Ala
1805						1810					1815			
Gln	Leu	Ala	Pro	Pro	Ser	Ala	Ala	Ser	Ala	Phe	Val	Gly	Ala	Gly
1820						1825					1830			
Ile	Ala	Gly	Ala	Ala	Val	Gly	Thr	Ile	Gly	Leu	Gly	Lys	Val	Leu
1835						1840					1845			
Val	Asp	Ile	Leu	Ala	Gly	Tyr	Gly	Ala	Gly	Val	Ala	Gly	Ala	Leu
1850						1855					1860			
Val	Ala	Phe	Lys	Ile	Met	Ser	Gly	Glu	Met	Pro	Ser	Ala	Glu	Asp
1865						1870					1875			
Met	Val	Asn	Leu	Leu	Pro	Ala	Ile	Leu	Ser	Pro	Gly	Ala	Leu	Val
1880						1885					1890			
Val	Gly	Ile	Val	Cys	Ala	Ala	Ile	Leu	Arg	Arg	His	Val	Gly	Pro
1895						1900					1905			

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Gly	Glu 1910	Gly	Ala	Val	Gln	Trp 1915	Met	Asn	Arg	Leu	Ile 1920	Ala	Phe	Ala
Ser	Arg 1925	Gly	Asn	His	Val	Ser 1930	Pro	Arg	His	Tyr	Val 1935	Pro	Glu	Ser
Glu	Pro 1940	Ala	Ala	Arg	Val	Thr 1945	Gln	Ile	Leu	Ser	Ser 1950	Leu	Thr	Ile
Thr	Gln 1955	Leu	Leu	Lys	Arg	Leu 1960	His	Gln	Trp	Ile	Asn 1965	Glu	Asp	Cys
Ser	Thr 1970	Pro	Cys	Ser	Ser	Ser 1975	Trp	Leu	Arg	Glu	Ile 1980	Trp	Asp	Trp
Ile	Cys 1985	Thr	Val	Leu	Thr	Asp 1990	Phe	Lys	Thr	Trp	Leu 1995	Gln	Ser	Lys
Leu	Leu 2000	Pro	Arg	Leu	Pro	Gly 2005	Val	Pro	Phe	Phe	Ser 2010	Cys	Gln	Arg
Gly	Tyr 2015	Lys	Gly	Val	Trp	Arg 2020	Gly	Asp	Gly	Ile	Met 2025	His	Thr	Thr
Cys	Pro 2030	Cys	Gly	Ala	Gln	Ile 2035	Thr	Gly	His	Val	Lys 2040	Asn	Gly	Ser
Met	Arg 2045	Ile	Val	Gly	Pro	Lys 2050	Thr	Cys	Ser	Asn	Thr 2055	Trp	Tyr	Gly
Thr	Phe 2060	Pro	Ile	Asn	Ala	Tyr 2065	Thr	Thr	Gly	Pro	Cys 2070	Thr	Pro	Ser
Pro	Ala 2075	Pro	Asn	Tyr	Ser	Lys 2080	Ala	Leu	Trp	Arg	Val 2085	Ala	Ala	Glu
Glu	Tyr 2090	Val	Glu	Val	Thr	Arg 2095	Val	Gly	Asp	Phe	His 2100	Tyr	Val	Thr
Gly	Met 2105	Thr	Thr	Asp	Asn	Val 2110	Lys	Cys	Pro	Cys	Gln 2115	Val	Pro	Ala
Pro	Glu 2120	Phe	Phe	Thr	Glu	Val 2125	Asp	Gly	Val	Arg	Leu 2130	His	Arg	Tyr
Ala	Pro 2135	Ala	Cys	Arg	Pro	Leu 2140	Leu	Arg	Glu	Glu	Val 2145	Val	Phe	Gln
Val	Gly 2150	Leu	His	Gln	Tyr	Leu 2155	Val	Gly	Ser	Gln	Leu 2160	Pro	Cys	Glu

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Pro	Glu	Pro	Asp	Val	Ala	Val	Leu	Thr	Ser	Met	Leu	Thr	Asp	Pro
	2165					2170					2175			
Ser	His	Ile	Thr	Ala	Glu	Thr	Ala	Lys	Arg	Arg	Leu	Ala	Arg	Gly
	2180					2185					2190			
Ser	Pro	Pro	Ser	Leu	Ala	Ser	Ser	Ser	Ala	Ser	Gln	Leu	Ser	Ala
	2195					2200					2205			
Pro	Ser	Leu	Lys	Ala	Thr	Cys	Thr	Thr	His	His	Asp	Ser	Pro	Asp
	2210					2215					2220			
Ala	Asp	Leu	Ile	Glu	Ala	Asn	Leu	Leu	Trp	Arg	Gln	Glu	Met	Gly
	2225					2230					2235			
Gly	Asn	Ile	Thr	Arg	Val	Glu	Ser	Glu	Asn	Lys	Val	Val	Ile	Leu
	2240					2245					2250			
Asp	Ser	Phe	Asp	Pro	Leu	Arg	Ala	Glu	Asp	Asp	Glu	Gly	Glu	Ile
	2255					2260					2265			
Ser	Val	Pro	Ala	Glu	Ile	Leu	Arg	Lys	Ser	Arg	Lys	Phe	Pro	Pro
	2270					2275					2280			
Ala	Leu	Pro	Ile	Trp	Ala	Pro	Pro	Asp	Tyr	Asn	Pro	Pro	Leu	Leu
	2285					2290					2295			
Glu	Ser	Trp	Lys	Asp	Pro	Asp	Tyr	Val	Pro	Pro	Val	Val	His	Gly
	2300					2305					2310			
Cys	Pro	Leu	Pro	Pro	Thr	Lys	Ala	Pro	Pro	Ile	Pro	Pro	Pro	Arg
	2315					2320					2325			
Arg	Lys	Arg	Thr	Val	Val	Leu	Thr	Glu	Ser	Thr	Val	Ser	Ser	Ala
	2330					2335					2340			
Leu	Ala	Glu	Leu	Ala	Thr	Lys	Thr	Phe	Gly	Ser	Ser	Gly	Ser	Ser
	2345					2350					2355			
Ala	Ile	Asp	Ser	Gly	Thr	Ala	Thr	Ala	Pro	Pro	Asp	Gln	Ala	Ser
	2360					2365					2370			
Gly	Asp	Gly	Asp	Arg	Glu	Ser	Asp	Val	Glu	Ser	Phe	Ser	Ser	Met
	2375					2380					2385			
Pro	Pro	Leu	Glu	Gly	Glu	Pro	Gly	Asp	Pro	Asp	Leu	Ser	Asp	Gly
	2390					2395					2400			
Ser	Trp	Ser	Thr	Val	Ser	Glu	Glu	Ala	Ser	Glu	Asp	Val	Val	Cys
	2405					2410					2415			

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Cys	Ser	Met	Ser	Tyr	Thr	Trp	Thr	Gly	Ala	Leu	Ile	Thr	Pro	Cys
	2420					2425					2430			
Ala	Ala	Glu	Glu	Ser	Lys	Leu	Pro	Ile	Asn	Pro	Leu	Ser	Asn	Ser
	2435					2440					2445			
Leu	Leu	Arg	His	His	Asn	Met	Val	Tyr	Ala	Thr	Thr	Ser	Arg	Ser
	2450					2455					2460			
Ala	Gly	Leu	Arg	Gln	Lys	Lys	Val	Thr	Phe	Asp	Arg	Leu	Gln	Val
	2465					2470					2475			
Leu	Asp	Asp	His	Tyr	Arg	Asp	Val	Leu	Lys	Glu	Met	Lys	Ala	Lys
	2480					2485					2490			
Ala	Ser	Thr	Val	Lys	Ala	Lys	Leu	Leu	Ser	Val	Glu	Glu	Ala	Cys
	2495					2500					2505			
Lys	Leu	Thr	Pro	Pro	His	Ser	Ala	Lys	Ser	Lys	Phe	Gly	Tyr	Gly
	2510					2515					2520			
Ala	Lys	Asp	Val	Arg	Ser	Leu	Ser	Ser	Arg	Ala	Val	Thr	His	Ile
	2525					2530					2535			
Arg	Ser	Val	Trp	Lys	Asp	Leu	Leu	Glu	Asp	Thr	Glu	Thr	Pro	Ile
	2540					2545					2550			
Ser	Thr	Thr	Ile	Met	Ala	Lys	Asn	Glu	Val	Phe	Cys	Val	Gln	Pro
	2555					2560					2565			
Glu	Lys	Gly	Gly	Arg	Lys	Pro	Ala	Arg	Leu	Ile	Val	Phe	Pro	Asp
	2570					2575					2580			
Leu	Gly	Val	Arg	Val	Cys	Glu	Lys	Met	Ala	Leu	Tyr	Asp	Val	Val
	2585					2590					2595			
Ser	Thr	Leu	Pro	Gln	Ala	Val	Met	Gly	Ser	Ser	Tyr	Gly	Phe	Gln
	2600					2605					2610			
Tyr	Ser	Pro	Lys	Gln	Arg	Val	Glu	Phe	Leu	Val	Asn	Thr	Trp	Lys
	2615					2620					2625			
Ser	Lys	Lys	Cys	Pro	Met	Gly	Phe	Ser	Tyr	Asp	Thr	Arg	Cys	Phe
	2630					2635					2640			
Asp	Ser	Thr	Val	Thr	Glu	Asn	Asp	Ile	Arg	Val	Glu	Glu	Ser	Ile
	2645					2650					2655			
Tyr	Gln	Cys	Cys	Asp	Leu	Ala	Pro	Glu	Ala	Lys	Leu	Ala	Ile	Lys
	2660					2665					2670			

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Ser	Leu 2675	Thr	Glu	Arg	Leu	Tyr 2680	Ile	Gly	Gly	Pro	Leu 2685	Thr	Asn	Ser
Lys	Gly 2690	Gln	Asn	Cys	Gly	Tyr 2695	Arg	Arg	Cys	Arg	Ala 2700	Ser	Gly	Val
Leu	Thr 2705	Thr	Ser	Cys	Gly	Asn 2710	Thr	Leu	Thr	Cys	Tyr 2715	Leu	Lys	Ala
Thr	Ala 2720	Ala	Cys	Arg	Ala	Ala 2725	Lys	Leu	Arg	Asp	Cys 2730	Thr	Met	Leu
Val	Asn 2735	Gly	Asp	Asp	Leu	Val 2740	Val	Ile	Cys	Glu	Ser 2745	Ala	Gly	Thr
Gln	Glu 2750	Asp	Ala	Ala	Ser	Leu 2755	Arg	Val	Phe	Thr	Glu 2760	Ala	Met	Thr
Arg	Tyr 2765	Ser	Ala	Pro	Pro	Gly 2770	Asp	Pro	Pro	Gln	Pro 2775	Glu	Tyr	Asp
Leu	Glu 2780	Leu	Ile	Thr	Ser	Cys 2785	Ser	Ser	Asn	Val	Ser 2790	Val	Ala	His
Asp	Ala 2795	Ser	Gly	Lys	Arg	Val 2800	Tyr	Tyr	Leu	Thr	Arg 2805	Asp	Pro	Thr
Thr	Pro 2810	Leu	Ala	Arg	Ala	Ala 2815	Trp	Glu	Thr	Ala	Arg 2820	His	Thr	Pro
Val	Asn 2825	Ser	Trp	Leu	Gly	Asn 2830	Ile	Ile	Met	Tyr	Ala 2835	Pro	Thr	Leu
Trp	Ala 2840	Arg	Met	Ile	Leu	Met 2845	Thr	His	Phe	Phe	Ser 2850	Ile	Leu	Leu
Ala	Gln 2855	Glu	Gln	Leu	Glu	Lys 2860	Thr	Leu	Asp	Cys	Gln 2865	Ile	Tyr	Gly
Ala	Cys 2870	Tyr	Ser	Ile	Glu	Pro 2875	Leu	Asp	Leu	Pro	Gln 2880	Ile	Ile	Glu
Arg	Leu 2885	His	Gly	Leu	Ser	Ala 2890	Phe	Ser	Leu	His	Ser 2895	Tyr	Ser	Pro
Gly	Glu 2900	Ile	Asn	Arg	Val	Ala 2905	Ser	Cys	Leu	Arg	Lys 2910	Leu	Gly	Val
Pro	Pro 2915	Leu	Arg	Ala	Trp	Arg 2920	His	Arg	Ala	Arg	Ser 2925	Val	Arg	Ala

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Lys Leu Leu Ser Gln Gly Gly Arg Ala Ala Thr Cys Gly Lys Tyr
2930 2935 2940

Leu Phe Asn Trp Ala Val Arg Thr Lys Leu Lys Leu Thr Pro Ile
2945 2950 2955

Pro Ala Ala Ser Arg Leu Asp Leu Ser Gly Trp Phe Val Ala Gly
2960 2965 2970

Tyr Ser Gly Gly Asp Ile Tyr His Ser Leu Ser Arg Ala Arg Pro
2975 2980 2985

Arg Trp Phe Met Leu Cys Leu Leu Leu Leu Ser Val Gly Val Gly
2990 2995 3000

Ile Tyr Leu Leu Pro Asn Arg
3005 3010

<210> 4
<211> 486
<212> PRT
<213> influenza A virus

<400> 4

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Gly Glu Arg Gln Asn Ala Thr Glu Ile Arg Ala Ser Val Gly Arg Met
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Val Gly Gly Ile Gly Arg Phe Tyr Ile Gln Met Cys Thr Glu Leu Lys
35 40 45

Leu Ser Asp His Glu Gly Arg Leu Ile Gln Asn Ser Ile Thr Ile Glu
50 55 60

Arg Met Val Leu Ser Ala Phe Asp Glu Arg Arg Asn Lys Tyr Leu Glu
65 70 75 80

Glu His Pro Ser Ala Gly Lys Asp Pro Lys Lys Thr Gly Gly Pro Ile
85 90 95

Tyr Arg Arg Arg Asp Gly Lys Trp Met Arg Glu Leu Ile Leu Tyr Asp
100 105 110

Lys Glu Glu Ile Arg Arg Ile Trp Arg Gln Ala Asn Asn Gly Glu Asp
115 120 125

Ala Thr Ala Gly Leu Thr His Met Met Ile Trp His Ser Asn Leu Asn
130 135 140

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Asp 145	Ala	Thr	Tyr	Gln	Arg 150	Thr	Arg	Ala	Leu	Val 155	Arg	Thr	Gly	Met	Asp 160
Pro	Arg	Met	Cys	Ser 165	Leu	Met	Gln	Gly	Ser 170	Thr	Leu	Pro	Arg	Arg	Ser
Gly	Ala	Ala	Gly 180	Ala	Ala	Val	Lys	Gly 185	Val	Gly	Thr	Met	Val 190	Met	Glu
Leu	Ile	Arg 195	Met	Ile	Lys	Arg	Gly 200	Ile	Asn	Asp	Arg	Asn 205	Phe	Trp	Arg
Gly	Glu 210	Asn	Gly	Arg	Arg	Thr 215	Arg	Ile	Ala	Tyr	Glu 220	Arg	Met	Cys	Asn
Ile 225	Leu	Lys	Gly	Lys	Phe 230	Gln	Thr	Ala	Ala	Gln 235	Arg	Ala	Met	Met	Asp 240
Gln	Val	Arg	Glu	Ser 245	Arg	Asn	Pro	Gly	Asn 250	Ala	Glu	Ile	Glu	Asp 255	Leu
Ile	Phe	Leu	Ala 260	Arg	Ser	Ala	Leu	Ile 265	Leu	Arg	Gly	Ser	Val 270	Ala	His
Lys	Ser	Cys 275	Leu	Pro	Ala	Cys	Val 280	Tyr	Gly	Leu	Ala	Val 285	Ala	Ser	Gly
Tyr	Asp 290	Phe	Glu	Arg	Glu	Gly 295	Tyr	Ser	Leu	Val	Gly 300	Ile	Asp	Pro	Phe
Arg 305	Leu	Leu	Gln	Asn	Ser 310	Gln	Val	Phe	Ser	Leu 315	Ile	Arg	Pro	Asn	Glu 320
Asn	Pro	Ala	His	Lys 325	Ser	Gln	Leu	Val	Trp 330	Met	Ala	Cys	His	Ser 335	Ala
Ala	Phe	Glu	Asp 340	Leu	Arg	Val	Ser	Ser 345	Phe	Ile	Arg	Gly	Thr 350	Arg	Val
Val	Pro	Arg 355	Gly	Gln	Leu	Ser	Thr 360	Arg	Gly	Val	Gln	Ile 365	Ala	Ser	Asn
Glu	Asn 370	Met	Glu	Thr	Met	Asp 375	Ser	Ser	Thr	Leu	Glu 380	Leu	Arg	Ser	Arg
Tyr 385	Trp	Ala	Ile	Arg	Thr 390	Arg	Ser	Gly	Gly	Asn 395	Thr	Asn	Gln	Gln	Arg 400
Ala	Ser	Ala	Gly	Gln 405	Ile	Ser	Val	Gln	Pro 410	Thr	Phe	Ser	Val	Gln 415	Arg

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Asn Leu Pro Phe Glu Arg Ala Thr Ile Met Ala Ala Phe Thr Gly Asn
420 425 430

Thr Glu Gly Arg Thr Ser Asp Met Arg Thr Glu Ile Ile Arg Met Met
435 440 445

Glu Asn Ala Arg Pro Glu Asp Val Ser Phe Gln Gly Arg Gly Val Phe
450 455 460

Glu Leu Ser Asp Glu Lys Ala Thr Asn Pro Ile Val Pro Ser Phe Asp
465 470 475 480

Met Ser Asn Glu Gly Ser
485

<210> 5
<211> 97
<212> PRT
<213> influenza A virus

<400> 5

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Cys Arg Cys Asn Asp Ser Ser Asp Pro Leu Val Val Ala Ala Ser Ile
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Ile Gly Ile Leu His Leu Ile Leu Trp Ile Leu Asp Arg Leu Phe Phe
35 40 45

Lys Cys Val Tyr Arg Leu Phe Lys His Gly Leu Lys Arg Gly Pro Ser
50 55 60

Thr Glu Gly Val Pro Glu Ser Met Arg Glu Glu Tyr Arg Lys Glu Gln
65 70 75 80

Gln Asn Ala Val Asp Ala Asp Asp Ser His Phe Val Ser Ile Glu Leu
85 90 95

Glu

<210> 6
<211> 498
<212> PRT
<213> influenza A virus

<400> 6

Met Ala Ser Gln Gly Thr Lys Arg Ser Tyr Glu Gln Met Glu Thr Asp
1 5 10 15

Gly Asp Arg Gln Asn Ala Thr Glu Ile Arg Ala Ser Val Gly Lys Met
20 25 30

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Ile Asp Gly Ile Gly Arg Phe Tyr Ile Gln Met Cys Thr Glu Leu Lys
35 40 45
Leu Ser Asp His Glu Gly Arg Leu Ile Gln Asn Ser Leu Thr Ile Glu
50 55 60
Lys Met Val Leu Ser Ala Phe Asp Glu Arg Arg Asn Lys Tyr Leu Glu
65 70 75 80
Glu His Pro Ser Ala Gly Lys Asp Pro Lys Lys Thr Gly Gly Pro Ile
85 90 95
Tyr Arg Arg Val Asp Gly Lys Trp Met Arg Glu Leu Val Leu Tyr Asp
100 105 110
Lys Glu Glu Ile Arg Arg Ile Trp Arg Gln Ala Asn Asn Gly Glu Asp
115 120 125
Ala Thr Ala Gly Leu Thr His Ile Met Ile Trp His Ser Asn Leu Asn
130 135 140
Asp Ala Thr Tyr Gln Arg Thr Arg Ala Leu Val Arg Thr Gly Met Asp
145 150 155 160
Pro Arg Met Cys Ser Leu Met Gln Gly Ser Thr Leu Pro Arg Arg Ser
165 170 175
Gly Ala Ala Gly Ala Ala Val Lys Gly Ile Gly Thr Met Val Met Glu
180 185 190
Leu Ile Arg Met Val Lys Arg Gly Ile Asn Asp Arg Asn Phe Trp Arg
195 200 205
Gly Glu Asn Gly Arg Lys Thr Arg Ser Ala Tyr Glu Arg Met Cys Asn
210 215 220
Ile Leu Lys Gly Lys Phe Gln Thr Ala Ala Gln Arg Ala Met Val Asp
225 230 235 240
Gln Val Arg Glu Ser Arg Asn Pro Gly Asn Ala Glu Ile Glu Asp Leu
245 250 255
Ile Phe Leu Ala Arg Ser Ala Leu Ile Leu Arg Gly Ser Val Ala His
260 265 270
Lys Ser Cys Leu Pro Ala Cys Ala Tyr Gly Pro Ala Val Ser Ser Gly
275 280 285
Tyr Asp Phe Glu Lys Glu Gly Tyr Ser Leu Val Gly Ile Asp Pro Phe
290 295 300

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Lys Leu Leu Gln Asn Ser Gln Ile Tyr Ser Leu Ile Arg Pro Asn Glu
305 310 315 320

Asn Pro Ala His Lys Ser Gln Leu Val Trp Met Ala Cys His Ser Ala
325 330 335

Ala Phe Glu Asp Leu Arg Leu Leu Ser Phe Ile Arg Gly Thr Lys Val
340 345 350

Ser Pro Arg Gly Lys Leu Ser Thr Arg Gly Val Gln Ile Ala Ser Asn
355 360 365

Glu Asn Met Asp Asn Met Gly Ser Ser Thr Leu Glu Leu Arg Ser Gly
370 375 380

Tyr Trp Ala Ile Arg Thr Arg Ser Gly Gly Asn Thr Asn Gln Gln Arg
385 390 395 400

Ala Ser Ala Gly Gln Thr Ser Val Gln Pro Thr Phe Ser Val Gln Arg
405 410 415

Asn Leu Pro Phe Glu Lys Ser Thr Ile Met Ala Ala Phe Thr Gly Asn
420 425 430

Thr Glu Gly Arg Thr Ser Asp Met Arg Ala Glu Ile Ile Arg Met Met
435 440 445

Glu Gly Ala Lys Pro Glu Glu Val Ser Phe Arg Gly Arg Gly Val Phe
450 455 460

Glu Leu Ser Asp Glu Lys Ala Thr Asn Pro Ile Val Pro Ser Phe Asp
465 470 475 480

Met Ser Asn Glu Gly Ser Tyr Phe Phe Gly Asp Asn Ala Glu Glu Tyr
485 490 495

Asp Asn

<210> 7
<211> 97
<212> PRT
<213> influenza A virus

<400> 7

Met Ser Leu Leu Thr Glu Val Glu Thr Pro Ile Arg Asn Glu Trp Gly
1 5 10 15

Cys Arg Cys Asn Gly Ser Ser Asp Pro Leu Ala Ile Ala Ala Asn Ile
20 25 30

pctdk2011050460-seq1

Ile Gly Ile Leu His Leu Ile Leu Trp Ile Leu Asp Arg Leu Phe Phe
35 40 45

Lys Cys Ile Tyr Arg Arg Phe Lys Tyr Gly Leu Lys Gly Gly Pro Ser
50 55 60

Thr Glu Gly Val Pro Lys Ser Met Arg Glu Glu Tyr Arg Lys Glu Gln
65 70 75 80

Gln Ser Ala Val Asp Ala Asp Asp Gly His Phe Val Ser Ile Glu Leu
85 90 95

Glu

<210> 8
<211> 498
<212> PRT
<213> influenza A virus

<400> 8

Met Ala Ser Gln Gly Thr Lys Arg Ser Tyr Glu Gln Met Glu Thr Asp
1 5 10 15

Gly Glu Arg Gln Asn Ala Thr Glu Ile Arg Ala Ser Val Gly Lys Met
20 25 30

Ile Gly Gly Ile Gly Arg Phe Tyr Ile Gln Met Cys Thr Glu Leu Lys
35 40 45

Leu Ser Asp Tyr Glu Gly Arg Leu Ile Gln Asn Ser Leu Thr Ile Glu
50 55 60

Arg Met Val Leu Ser Ala Phe Asp Glu Arg Arg Asn Lys Tyr Leu Glu
65 70 75 80

Glu His Pro Ser Ala Gly Lys Asp Pro Lys Lys Thr Gly Gly Pro Ile
85 90 95

Tyr Arg Arg Val Asn Gly Lys Trp Met Arg Glu Leu Ile Leu Tyr Asp
100 105 110

Lys Glu Glu Ile Arg Arg Ile Trp Arg Gln Ala Asn Asn Gly Asp Asp
115 120 125

Ala Thr Ala Gly Leu Thr His Met Met Ile Trp His Ser Asn Leu Asn
130 135 140

Asp Ala Thr Tyr Gln Arg Thr Arg Ala Leu Val Arg Thr Gly Met Asp
145 150 155 160

pctdk2011050460-seq1

Pro Arg Met Cys Ser₁₆₅ Leu Met Gln Gly Ser₁₇₀ Thr Leu Pro Arg Arg₁₇₅ Ser
Gly Ala Ala Gly₁₈₀ Ala Ala Val Lys Gly₁₈₅ Val Gly Thr Met Val₁₉₀ Met Glu
Leu Val Arg₁₉₅ Met Ile Lys Arg Gly₂₀₀ Ile Asn Asp Arg Asn₂₀₅ Phe Trp Arg
Gly Glu₂₁₀ Asn Gly Arg Lys Thr₂₁₅ Arg Ile Ala Tyr Glu₂₂₀ Arg Met Cys Asn
Ile₂₂₅ Leu Lys Gly Lys Phe₂₃₀ Gln Thr Ala Ala Gln₂₃₅ Lys Ala Met Met Asp₂₄₀
Gln Val Arg Glu Ser₂₄₅ Arg Asp Pro Gly Asn₂₅₀ Ala Glu Phe Glu Asp₂₅₅ Leu
Thr Phe Leu Ala₂₆₀ Arg Ser Ala Leu Ile₂₆₅ Leu Arg Gly Ser Val₂₇₀ Ala His
Lys Ser Cys₂₇₅ Leu Pro Ala Cys Val₂₈₀ Tyr Gly Pro Ala Val₂₈₅ Ala Ser Gly
Tyr Asp₂₉₀ Phe Glu Arg Glu Gly₂₉₅ Tyr Ser Leu Val Gly₃₀₀ Ile Asp Pro Phe
Arg₃₀₅ Leu Leu Gln Asn Ser₃₁₀ Gln Val Tyr Ser Leu₃₁₅ Ile Arg Pro Asn Glu₃₂₀
Asn Pro Ala His Lys₃₂₅ Ser Gln Leu Val Trp₃₃₀ Met Ala Cys His Ser₃₃₅ Ala
Ala Phe Glu Asp₃₄₀ Leu Arg Val Leu Ser₃₄₅ Phe Ile Lys Gly Thr₃₅₀ Lys Val
Val Pro Arg₃₅₅ Gly Lys Leu Ser Thr₃₆₀ Arg Gly Val Gln Ile₃₆₅ Ala Ser Asn
Glu Asn₃₇₀ Met Glu Thr Met Glu₃₇₅ Ser Ser Thr Leu Glu₃₈₀ Leu Arg Ser Arg
Tyr Trp Ala Ile Arg Thr₃₉₀ Arg Ser Gly Gly Asn₃₉₅ Thr Asn Gln Gln Arg₄₀₀
Ala Ser Ala Gly Gln₄₀₅ Ile Ser Ile Gln Pro Thr Phe Ser Val Gln₄₁₅ Arg
Asn Leu Pro Phe₄₂₀ Asp Arg Thr Thr Val₄₂₅ Met Ala Ala Phe Thr₄₃₀ Gly Asn

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Thr Glu Gly Arg Thr Ser Asp Met Arg Thr Glu Ile Ile Arg Met Met
435 440 445

Glu Ser Ala Arg Pro Glu Asp Val Ser Phe Gln Gly Arg Gly Val Phe
450 455 460

Glu Leu Ser Asp Glu Lys Ala Ala Ser Pro Ile Val Pro Ser Phe Asp
465 470 475 480

Met Ser Asn Glu Gly Ser Tyr Phe Phe Gly Asp Asn Ala Glu Glu Tyr
485 490 495

Asp Asn

<210> 9
<211> 97
<212> PRT
<213> influenza A virus

<400> 9

Met Ser Leu Leu Thr Glu Val Glu Thr Pro Ile Arg Asn Glu Trp Gly
1 5 10 15

Cys Arg Cys Asn Asp Ser Ser Asp Pro Leu Val Val Ala Ala Ser Ile
20 25 30

Ile Gly Ile Leu His Phe Ile Leu Trp Ile Leu Asp Arg Leu Phe Phe
35 40 45

Lys Cys Ile Tyr Arg Phe Phe Lys His Gly Leu Lys Arg Gly Pro Ser
50 55 60

Thr Glu Gly Val Pro Glu Ser Met Arg Glu Glu Tyr Arg Lys Glu Gln
65 70 75 80

Gln Ser Ala Val Asp Ala Asp Asp Ser His Phe Val Ser Ile Glu Leu
85 90 95

Glu

<210> 10
<211> 498
<212> PRT
<213> influenza A virus

<400> 10

Met Ala Ser Gln Gly Thr Lys Arg Ser Tyr Glu Gln Met Glu Thr Asp
1 5 10 15

Gly Glu Arg Gln Asn Ala Thr Glu Ile Arg Ala Ser Val Gly Lys Met
20 25 30

pctdk2011050460-seq1

Ile Asp Gly Ile Gly Arg Phe Tyr Ile Gln Met Cys Thr Glu Leu Lys
35 40 45

Leu Ser Asp Tyr Glu Gly Arg Leu Ile Gln Asn Ser Leu Thr Ile Glu
50 55 60

Arg Met Val Leu Ser Ala Phe Asp Glu Arg Arg Asn Lys Tyr Leu Glu
65 70 75 80

Glu His Pro Ser Ala Gly Lys Asp Pro Lys Lys Thr Gly Gly Pro Ile
85 90 95

Tyr Lys Arg Val Asp Gly Lys Trp Met Arg Glu Leu Val Leu Tyr Asp
100 105 110

Lys Glu Glu Ile Arg Arg Ile Trp Arg Gln Ala Asn Asn Gly Asp Asp
115 120 125

Ala Thr Ala Gly Leu Thr His Met Met Ile Trp His Ser Asn Leu Asn
130 135 140

Asp Thr Thr Tyr Gln Arg Thr Arg Ala Leu Val Arg Thr Gly Met Asp
145 150 155 160

Pro Arg Met Cys Ser Leu Met Gln Gly Ser Thr Leu Pro Arg Arg Ser
165 170 175

Gly Ala Ala Gly Ala Ala Val Lys Gly Val Gly Thr Met Val Met Glu
180 185 190

Leu Ile Arg Met Ile Lys Arg Gly Ile Asn Asp Arg Asn Phe Trp Arg
195 200 205

Gly Glu Asn Gly Arg Lys Thr Arg Ser Ala Tyr Glu Arg Met Cys Asn
210 215 220

Ile Leu Lys Gly Lys Phe Gln Thr Ala Ala Gln Arg Ala Met Met Asp
225 230 235 240

Gln Val Arg Glu Ser Arg Asn Pro Gly Asn Ala Glu Ile Glu Asp Leu
245 250 255

Ile Phe Leu Ala Arg Ser Ala Leu Ile Leu Arg Gly Ser Val Ala His
260 265 270

Lys Ser Cys Leu Pro Ala Cys Val Tyr Gly Pro Ala Ile Ala Ser Gly
275 280 285

Tyr Asn Phe Glu Lys Glu Gly Tyr Ser Leu Val Gly Ile Asp Pro Phe
290 295 300

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Lys Leu Leu Gln Asn Ser Gln Val Tyr Ser Leu Ile Arg Pro Asn Glu
305 310 315 320

Asn Pro Ala His Lys Ser Gln Leu Val Trp Met Ala Cys Asn Ser Ala
325 330 335

Ala Phe Glu Asp Leu Arg Val Leu Ser Phe Ile Arg Gly Thr Lys Val
340 345 350

Ser Pro Arg Gly Lys Leu Ser Thr Arg Gly Val Gln Ile Ala Ser Asn
355 360 365

Glu Asn Met Asp Thr Met Glu Ser Ser Thr Leu Glu Leu Arg Ser Arg
370 375 380

Tyr Trp Ala Ile Arg Thr Arg Ser Gly Gly Asn Thr Asn Gln Gln Arg
385 390 395 400

Ala Ser Ala Gly Gln Ile Ser Val Gln Pro Ala Phe Ser Val Gln Arg
405 410 415

Asn Leu Pro Phe Asp Lys Pro Thr Ile Met Ala Ala Phe Thr Gly Asn
420 425 430

Thr Glu Gly Arg Thr Ser Asp Met Arg Ala Glu Ile Ile Arg Met Met
435 440 445

Glu Gly Ala Lys Pro Glu Glu Met Ser Phe Gln Gly Arg Gly Val Phe
450 455 460

Glu Leu Ser Asp Glu Lys Ala Thr Asn Pro Ile Val Pro Ser Phe Asp
465 470 475 480

Met Ser Asn Glu Gly Ser Tyr Phe Phe Gly Asp Asn Ala Glu Glu Tyr
485 490 495

Asp Asn

<210> 11
<211> 551
<212> PRT
<213> Human herpesvirus 5

<400> 11

Met Ala Ser Val Leu Gly Pro Ile Ser Gly His Val Leu Lys Ala Val
1 5 10 15

Phe Ser Arg Gly Asp Thr Pro Val Leu Pro His Glu Thr Arg Leu Leu
20 25 30

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Gln Thr Gly Ile His Val Arg Val Ser Gln Pro Ser Leu Ile Leu Val
35 40 45
Ser Gln Tyr Thr Pro Asp Ser Thr Pro Cys His Arg Gly Asp Asn Gln
50 55 60
Leu Gln Val Gln His Thr Tyr Phe Thr Gly Ser Glu Val Glu Asn Val
65 70 75 80
Ser Val Asn Val His Asn Pro Thr Gly Arg Ser Ile Cys Pro Ser Gln
85 90 95
Glu Pro Met Ser Ile Tyr Val Tyr Ala Leu Pro Leu Lys Met Leu Asn
100 105 110
Ile Pro Ser Ile Asn Val His His Tyr Pro Ser Ala Ala Glu Arg Lys
115 120 125
His Arg His Leu Pro Val Ala Asp Ala Val Ile His Ala Ser Gly Lys
130 135 140
Gln Met Trp Gln Ala Arg Leu Thr Val Ser Gly Leu Ala Trp Thr Arg
145 150 155 160
Gln Gln Asn Gln Trp Lys Glu Pro Asp Val Tyr Tyr Thr Ser Ala Phe
165 170 175
Val Phe Pro Thr Lys Asp Val Ala Leu Arg His Val Val Cys Ala His
180 185 190
Glu Leu Val Cys Ser Met Glu Asn Thr Arg Ala Thr Lys Met Gln Val
195 200 205
Ile Gly Asp Gln Tyr Val Lys Val Tyr Leu Glu Ser Phe Cys Glu Asp
210 215 220
Val Pro Ser Gly Lys Leu Phe Met His Val Thr Leu Gly Ser Asp Val
225 230 235 240
Glu Glu Asp Leu Thr Met Thr Arg Asn Pro Gln Pro Phe Met Arg Pro
245 250 255
His Glu Arg Asn Gly Phe Thr Val Leu Cys Pro Lys Asn Met Ile Ile
260 265 270
Lys Pro Gly Lys Ile Ser His Ile Met Leu Asp Val Ala Phe Thr Ser
275 280 285
His Glu His Phe Gly Leu Leu Cys Pro Lys Ser Ile Pro Gly Leu Ser
290 295 300

pctdk2011050460-seq1

Ile Ser Gly Asn Leu Leu Met Asn Gly Gln Gln Ile Phe Leu Glu Val
305 310 315 320

Gln Ala Ile Arg Glu Thr Val Glu Leu Arg Gln Tyr Asp Pro Val Ala
325 330 335

Ala Leu Phe Phe Phe Asp Ile Asp Leu Leu Leu Gln Arg Gly Pro Gln
340 345 350

Tyr Ser Glu His Pro Thr Phe Thr Ser Gln Tyr Arg Ile Gln Gly Lys
355 360 365

Leu Glu Tyr Arg His Thr Trp Asp Arg His Asp Glu Gly Ala Ala Gln
370 375 380

Gly Asp Asp Asp Val Trp Thr Ser Gly Ser Asp Ser Asp Glu Glu Leu
385 390 395 400

Val Thr Thr Glu Arg Lys Thr Pro Arg Val Thr Gly Gly Gly Ala Met
405 410 415

Ala Gly Ala Ser Thr Ser Ala Gly Arg Lys Arg Lys Ser Ala Ser Ser
420 425 430

Ala Thr Ala Cys Thr Ala Gly Val Met Thr Arg Gly Arg Leu Lys Ala
435 440 445

Glu Ser Thr Val Ala Pro Glu Glu Asp Thr Asp Glu Asp Ser Asp Asn
450 455 460

Glu Ile His Asn Pro Ala Val Phe Thr Trp Pro Pro Trp Gln Ala Gly
465 470 475 480

Ile Leu Ala Arg Asn Leu Val Pro Met Val Ala Thr Val Gln Gly Gln
485 490 495

Asn Leu Lys Tyr Gln Glu Phe Phe Trp Asp Ala Asn Asp Ile Tyr Arg
500 505 510

Ile Phe Ala Glu Leu Glu Gly Val Trp Gln Pro Ala Ala Gln Pro Lys
515 520 525

Arg Arg Arg His Arg Gln Asp Ala Leu Pro Gly Pro Cys Ile Ala Ser
530 535 540

Thr Pro Lys Lys His Arg Gly
545 550

<210> 12
<211> 158
<212> PRT

<213> Human papillomavirus type 16

<400> 12

Met His Gln Lys Arg Thr Ala Met Phe Gln Asp Pro Gln Glu Arg Pro
 1 5 10 15

Gly Lys Leu Pro Gln Leu Cys Thr Glu Leu Gln Thr Thr Ile His Asp
 20 25 30

Ile Ile Leu Glu Cys Val Tyr Cys Lys Gln Gln Leu Leu Arg Arg Glu
 35 40 45

Val Tyr Asp Phe Ala Phe Arg Asp Leu Cys Ile Val Tyr Arg Asp Gly
 50 55 60

Asn Pro Tyr Ala Val Cys Asp Lys Cys Leu Lys Phe Tyr Ser Lys Ile
 65 70 75 80

Ser Glu Tyr Arg His Tyr Cys Tyr Ser Val Tyr Gly Thr Thr Leu Glu
 85 90 95

Gln Gln Tyr Asn Lys Pro Leu Cys Asp Leu Leu Ile Arg Cys Ile Asn
 100 105 110

Cys Gln Lys Pro Leu Cys Pro Glu Glu Lys Gln Arg His Leu Asp Lys
 115 120 125

Lys Gln Arg Phe His Asn Ile Arg Gly Arg Trp Thr Gly Arg Cys Met
 130 135 140

Ser Cys Cys Arg Ser Ser Arg Thr Arg Arg Glu Thr Gln Leu
 145 150 155

<210> 13

<211> 97

<212> PRT

<213> Influenza A virus

<400> 13

Met Ser Leu Leu Thr Glu Val Glu Thr Pro Ile Arg Asn Glu Trp Gly
 1 5 10 15

Cys Arg Cys Asn Gly Ser Ser Asp Pro Leu Ala Ile Ala Ala Asn Ile
 20 25 30

Ile Gly Ile Leu His Leu Ile Leu Trp Ile Leu Asp Arg Leu Phe Phe
 35 40 45

Lys Cys Ile Tyr Arg Arg Phe Lys Tyr Gly Leu Lys Gly Gly Pro Ser
 50 55 60

Thr Glu Gly Val Pro Lys Ser Met Arg Glu Glu Tyr Arg Lys Glu Gln

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65 70 75 80

Gln Ser Ala Val Asp Ala Asp Asp Gly His Phe Val Ser Ile Glu Leu

85 90 95

Glu

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<210> 14
<211> 759
<212> PRT
<213> Influenza A virus
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<400> 14

Met Glu Arg Ile Lys Glu Leu Arg Asn Leu Met Ser Gln Ser Arg Thr
1 5 10 15

Arg Glu Ile Leu Thr Lys Thr Thr Val Asp His Met Ala Ile Ile Lys
20 25 30

Lys Tyr Thr Ser Gly Arg Gln Glu Lys Asn Pro Ala Leu Arg Met Lys
35 40 45

Trp Met Met Ala Met Lys Tyr Pro Ile Thr Ala Asp Lys Arg Ile Thr
50 55 60

Glu Met Ile Pro Glu Arg Asn Glu Gln Gly Gln Thr Leu Trp Ser Lys
65 70 75 80

Met Asn Asp Ala Gly Ser Asp Arg Val Met Val Ser Pro Leu Ala Val
85 90 95

Thr Trp Trp Asn Arg Asn Gly Pro Met Thr Asn Thr Val His Tyr Pro
100 105 110

Lys Ile Tyr Lys Thr Tyr Phe Glu Arg Val Glu Arg Leu Lys His Gly
115 120 125

Thr Phe Gly Pro Val His Phe Arg Asn Gln Val Lys Ile Arg Arg Arg
130 135 140

Val Asp Ile Asn Pro Gly His Ala Asp Leu Ser Ala Lys Glu Ala Gln
145 150 155 160

Asp Val Ile Met Glu Val Val Phe Pro Asn Glu Val Gly Ala Arg Ile
165 170 175

Leu Thr Ser Glu Ser Gln Leu Thr Ile Thr Lys Glu Lys Lys Glu Glu
180 185 190

Leu Gln Asp Cys Lys Ile Ser Pro Leu Met Val Ala Tyr Met Leu Glu
195 200 205

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Arg Glu Leu Val Arg Lys Thr Arg Phe Leu Pro Val Ala Gly Gly Thr
210 215 220
Ser Ser Val Tyr Ile Glu Val Leu His Leu Thr Gln Gly Thr Cys Trp
225 230 235 240
Glu Gln Met Tyr Thr Pro Gly Gly Glu Val Lys Asn Asp Asp Val Asp
245 250 255
Gln Ser Leu Ile Ile Ala Ala Arg Asn Ile Val Arg Arg Ala Ala Val
260 265 270
Ser Ala Asp Pro Leu Ala Ser Leu Leu Glu Met Cys His Ser Thr Gln
275 280 285
Ile Gly Gly Ile Arg Met Val Asp Ile Leu Lys Gln Asn Pro Thr Glu
290 295 300
Glu Gln Ala Val Gly Ile Cys Lys Ala Ala Met Gly Leu Arg Ile Ser
305 310 315 320
Ser Ser Phe Ser Phe Gly Gly Phe Thr Phe Lys Arg Thr Ser Gly Ser
325 330 335
Ser Val Lys Arg Glu Glu Glu Val Leu Thr Gly Asn Leu Gln Thr Leu
340 345 350
Lys Ile Arg Val His Glu Gly Tyr Glu Glu Phe Thr Met Val Gly Arg
355 360 365
Arg Ala Thr Ala Ile Leu Arg Lys Ala Thr Arg Arg Leu Ile Gln Leu
370 375 380
Ile Val Ser Gly Arg Asp Glu Gln Ser Ile Ala Glu Ala Ile Ile Val
385 390 395 400
Ala Met Val Phe Ser Gln Glu Asp Cys Met Ile Lys Ala Val Arg Gly
405 410 415
Asp Leu Asn Phe Val Asn Arg Ala Asn Gln Arg Leu Asn Pro Met His
420 425 430
Gln Leu Leu Arg His Phe Gln Lys Asp Ala Lys Val Leu Phe Gln Asn
435 440 445
Trp Gly Val Glu Pro Ile Asp Asn Val Met Gly Met Ile Gly Ile Leu
450 455 460
Pro Asp Met Thr Pro Ser Ile Glu Met Ser Met Arg Gly Val Arg Ile
465 470 475 480

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Ser Lys Met Gly Val Asp Glu Tyr Ser Ser Thr Glu Arg Val Val Val
485 490 495

Ser Ile Asp Arg Phe Leu Arg Val Arg Asp Gln Arg Gly Asn Val Leu
500 505 510

Leu Ser Pro Glu Glu Val Ser Glu Thr Gln Gly Thr Glu Lys Leu Thr
515 520 525

Ile Thr Tyr Ser Ser Ser Met Met Trp Glu Ile Asn Gly Pro Glu Ser
530 535 540

Val Leu Val Asn Thr Tyr Gln Trp Ile Ile Arg Asn Trp Glu Thr Val
545 550 555 560

Lys Ile Gln Trp Ser Gln Asn Pro Thr Met Leu Tyr Asn Lys Met Glu
565 570 575

Phe Glu Pro Phe Gln Ser Leu Val Pro Lys Ala Ile Arg Gly Gln Tyr
580 585 590

Ser Gly Phe Val Arg Thr Leu Phe Gln Gln Met Arg Asp Val Leu Gly
595 600 605

Thr Phe Asp Thr Ala Gln Ile Ile Lys Leu Leu Pro Phe Ala Ala Ala
610 615 620

Pro Pro Lys Gln Ser Arg Met Gln Phe Ser Ser Phe Thr Val Asn Val
625 630 635 640

Arg Gly Ser Gly Met Arg Ile Leu Val Arg Gly Asn Ser Pro Val Phe
645 650 655

Asn Tyr Asn Lys Ala Thr Lys Arg Leu Thr Val Leu Gly Lys Asp Ala
660 665 670

Gly Thr Leu Thr Glu Asp Pro Asp Glu Gly Thr Ala Gly Val Glu Ser
675 680 685

Ala Val Leu Arg Gly Phe Leu Ile Leu Gly Lys Glu Asp Arg Arg Tyr
690 695 700

Gly Pro Ala Leu Ser Ile Asn Glu Leu Ser Asn Leu Ala Lys Gly Glu
705 710 715 720

Lys Ala Asn Val Leu Ile Gly Gln Gly Asp Val Val Leu Val Met Lys
725 730 735

Arg Lys Arg Asp Ser Ser Ile Leu Thr Asp Ser Gln Thr Ala Thr Lys
740 745 750

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Arg Ile Arg Met Ala Ile Asn
755

<210> 15
<211> 716
<212> PRT
<213> Influenza A virus

<400> 15

Met Glu Asp Phe Val Arg Gln Cys Phe Asn Pro Met Ile Val Glu Leu
1 5 10 15

Ala Glu Lys Thr Met Lys Glu Tyr Gly Glu Asp Leu Lys Ile Glu Thr
20 25 30

Asn Lys Phe Ala Ala Ile Cys Thr His Leu Glu Val Cys Phe Met Tyr
35 40 45

Ser Asp Phe His Phe Ile Asn Glu Gln Gly Glu Ser Ile Ile Val Glu
50 55 60

Leu Gly Asp Pro Asn Ala Leu Leu Lys His Arg Phe Glu Ile Ile Glu
65 70 75 80

Gly Arg Asp Arg Thr Met Ala Trp Thr Val Val Asn Ser Ile Cys Asn
85 90 95

Thr Thr Gly Ala Glu Lys Pro Lys Phe Leu Pro Asp Leu Tyr Asp Tyr
100 105 110

Lys Glu Asn Arg Phe Ile Glu Ile Gly Val Thr Arg Arg Glu Val His
115 120 125

Ile Tyr Tyr Leu Glu Lys Ala Asn Lys Ile Lys Ser Glu Lys Thr His
130 135 140

Ile His Ile Phe Ser Phe Thr Gly Glu Glu Met Ala Thr Lys Ala Asp
145 150 155 160

Tyr Thr Leu Asp Glu Glu Ser Arg Ala Arg Ile Lys Thr Arg Leu Phe
165 170 175

Thr Ile Arg Gln Glu Met Ala Ser Arg Gly Leu Trp Asp Ser Phe Arg
180 185 190

Gln Ser Glu Arg Gly Glu Glu Thr Ile Glu Glu Arg Phe Glu Ile Thr
195 200 205

Gly Thr Met Arg Lys Leu Ala Asp Gln Ser Leu Pro Pro Asn Phe Ser
210 215 220

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Ser Leu Glu Asn Phe Arg Ala Tyr Val Asp Gly Phe Glu Pro Asn Gly
225      230      235      240

Tyr Ile Glu Gly Lys Leu Ser Gln Met Ser Lys Glu Val Asn Ala Arg
      245      250      255

Ile Glu Pro Phe Leu Lys Thr Thr Pro Arg Pro Leu Arg Leu Pro Asn
      260      265      270

Gly Pro Pro Cys Ser Gln Arg Ser Lys Phe Leu Leu Met Asp Ala Leu
      275      280      285

Lys Leu Ser Ile Glu Asp Pro Ser His Glu Gly Glu Gly Ile Pro Leu
      290      295      300

Tyr Asp Ala Ile Lys Cys Met Arg Thr Phe Phe Gly Trp Lys Glu Pro
305      310      315      320

Asn Val Val Lys Pro His Glu Lys Gly Ile Asn Pro Asn Tyr Leu Leu
      325      330      335

Ser Trp Lys Gln Val Leu Ala Glu Leu Gln Asp Ile Glu Asn Glu Glu
      340      345      350

Lys Ile Pro Lys Thr Lys Asn Met Lys Lys Thr Ser Gln Leu Lys Trp
      355      360      365

Ala Leu Gly Glu Asn Met Ala Pro Glu Lys Val Asp Phe Asp Asp Cys
      370      375      380

Lys Asp Val Gly Asp Leu Lys Gln Tyr Asp Ser Asp Glu Pro Glu Leu
385      390      395      400

Arg Ser Leu Ala Ser Trp Ile Gln Asn Glu Phe Asn Lys Ala Cys Glu
      405      410      415

Leu Thr Asp Ser Ser Trp Ile Glu Leu Asp Glu Ile Gly Glu Asp Val
      420      425      430

Ala Pro Ile Glu His Ile Ala Ser Met Arg Arg Asn Tyr Phe Thr Ser
      435      440      445

Glu Val Ser His Cys Arg Ala Thr Glu Tyr Ile Met Lys Gly Val Tyr
      450      455      460

Ile Asn Thr Ala Leu Leu Asn Ala Ser Cys Ala Ala Met Asp Asp Phe
465      470      475      480

Gln Leu Ile Pro Met Ile Ser Lys Cys Arg Thr Lys Glu Gly Arg Arg
      485      490      495

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Lys Thr Asn Leu Tyr Gly Phe Ile Ile Lys Gly Arg Ser His Leu Arg
500 505 510

Asn Asp Thr Asp Val Val Asn Phe Val Ser Met Glu Phe Ser Leu Thr
515 520 525

Asp Pro Arg Leu Glu Pro His Lys Trp Glu Lys Tyr Cys Val Leu Glu
530 535 540

Ile Gly Asp Met Leu Leu Arg Ser Ala Ile Gly Gln Val Ser Arg Pro
545 550 555 560

Met Phe Leu Tyr Val Arg Thr Asn Gly Thr Ser Lys Ile Lys Met Lys
565 570 575

Trp Gly Met Glu Met Arg Arg Cys Leu Leu Gln Ser Leu Gln Gln Ile
580 585 590

Glu Ser Met Ile Glu Ala Glu Ser Ser Val Lys Glu Lys Asp Met Thr
595 600 605

Lys Glu Phe Phe Glu Asn Lys Ser Glu Thr Trp Pro Ile Gly Glu Ser
610 615 620

Pro Lys Gly Val Glu Glu Ser Ser Ile Gly Lys Val Cys Arg Thr Leu
625 630 635 640

Leu Ala Lys Ser Val Phe Asn Ser Leu Tyr Ala Ser Pro Gln Leu Glu
645 650 655

Gly Phe Ser Ala Glu Ser Arg Lys Leu Leu Leu Ile Val Gln Ala Leu
660 665 670

Arg Asp Asn Leu Glu Pro Gly Thr Phe Asp Leu Gly Gly Leu Tyr Glu
675 680 685

Ala Ile Glu Glu Cys Leu Ile Asn Asp Pro Trp Val Leu Leu Asn Ala
690 695 700

Ser Trp Phe Asn Ser Phe Leu Thr His Ala Leu Ser
705 710 715

<210> 16
<211> 230
<212> PRT
<213> Influenza A virus

<400> 16

Met Asp Pro Asn Thr Val Ser Ser Phe Gln Val Asp Cys Phe Leu Trp
1 5 10 15

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His Val Arg Lys Arg Val Ala Asp Gln Glu Leu Gly Asp Ala Pro Phe
20 25 30

Leu Asp Arg Leu Arg Arg Asp Gln Lys Ser Leu Arg Gly Arg Gly Ser
35 40 45

Thr Leu Gly Leu Asp Ile Glu Thr Ala Thr Arg Ala Gly Lys Gln Ile
50 55 60

Val Glu Arg Ile Leu Lys Glu Glu Ser Asp Glu Ala Leu Lys Met Thr
65 70 75 80

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

Glu Met Ser Arg Glu Trp Ser Met Leu Ile Pro Lys Gln Lys Val Ala
100 105 110

Gly Pro Leu Cys Ile Arg Met Asp Gln Ala Ile Met Asp Lys Asn Ile
115 120 125

Ile Leu Lys Ala Asn Phe Ser Val Ile Phe Asp Arg Leu Glu Thr Leu
130 135 140

Ile Leu Leu Arg Ala Phe Thr Glu Glu Gly Ala Ile Val Gly Glu Ile
145 150 155 160

Ser Pro Leu Pro Ser Leu Pro Gly His Thr Ala Glu Asp Val Lys Asn
165 170 175

Ala Val Gly Val Leu Ile Gly Gly Leu Glu Trp Asn Asp Asn Thr Val
180 185 190

Arg Val Ser Glu Thr Leu Gln Arg Phe Ala Trp Arg Ser Ser Asn Glu
195 200 205

Asn Gly Arg Pro Pro Leu Thr Pro Lys Gln Lys Arg Glu Met Ala Gly
210 215 220

Thr Ile Arg Ser Glu Val
225 230

<210> 17
<211> 121
<212> PRT
<213> Influenza A virus

<400> 17

Met Asp Pro Asn Thr Val Ser Ser Phe Gln Asp Ile Leu Leu Arg Met
1 5 10 15

Ser Lys Met Gln Leu Glu Ser Ser Ser Glu Asp Leu Asn Gly Met Ile
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20

25

30

Thr Gln Phe Glu Ser Leu Lys Leu Tyr Arg Asp Ser Leu Gly Glu Ala
 35 40 45

Val Met Arg Met Gly Asp Leu His Ser Leu Gln Asn Arg Asn Glu Lys
 50 55 60

Trp Arg Glu Gln Leu Gly Gln Lys Phe Glu Glu Ile Arg Trp Leu Ile
 65 70 75 80

Glu Glu Val Arg His Lys Leu Lys Val Thr Glu Asn Ser Phe Glu Gln
 85 90 95

Ile Thr Phe Met Gln Ala Leu His Leu Leu Leu Glu Val Glu Gln Glu
 100 105 110

Ile Arg Thr Phe Ser Phe Gln Leu Ile
 115 120

<210> 18

<211> 454

<212> PRT

<213> Influenza A virus

<400> 18

Met Asn Pro Asn Gln Lys Ile Ile Thr Ile Gly Ser Ile Cys Leu Val
 1 5 10 15

Val Gly Leu Ile Ser Leu Ile Leu Gln Ile Gly Asn Ile Ile Ser Ile
 20 25 30

Trp Ile Ser His Ser Ile Gln Thr Gly Ser Gln Asn His Thr Gly Ile
 35 40 45

Cys Asn Gln Asn Ile Ile Thr Tyr Lys Asn Ser Thr Trp Val Lys Asp
 50 55 60

Thr Thr Ser Val Ile Leu Thr Gly Asn Ser Ser Leu Cys Pro Ile Arg
 65 70 75 80

Gly Trp Ala Ile Tyr Ser Lys Asp Asn Ser Ile Arg Ile Gly Ser Lys
 85 90 95

Gly Asp Val Phe Val Ile Arg Glu Pro Phe Ile Ser Cys Ser His Leu
 100 105 110

Glu Cys Arg Thr Phe Phe Leu Thr Gln Gly Ala Leu Leu Asn Asp Arg
 115 120 125

His Ser Asn Gly Thr Val Lys Asp Arg Ser Pro Tyr Arg Ala Leu Met
 130 135 140

pctdk2011050460-seq1

Ser Cys Pro Val Gly Glu Ala Pro Ser Pro Tyr Asn Ser Arg Phe Glu
145 150 155 160

Ser Val Ala Trp Ser Ala Ser Ala Cys His Asp Gly Met Gly Trp Leu
165 170 175

Thr Ile Gly Ile Ser Gly Pro Asp Asn Gly Ala Val Ala Val Leu Lys
180 185 190

Tyr Asn Gly Ile Ile Thr Glu Thr Ile Lys Ser Trp Arg Lys Lys Ile
195 200 205

Leu Arg Thr Gln Glu Ser Glu Cys Ala Cys Val Asn Gly Ser Cys Phe
210 215 220

Thr Ile Met Thr Asp Gly Pro Ser Asp Gly Leu Ala Ser Tyr Lys Ile
225 230 235 240

Phe Lys Ile Glu Lys Gly Lys Val Thr Lys Ser Ile Glu Leu Asn Ala
245 250 255

Pro Asn Ser His Tyr Glu Glu Cys Ser Cys Tyr Pro Asp Thr Gly Lys
260 265 270

Val Met Cys Val Cys Arg Asp Asn Trp His Gly Ser Asn Arg Pro Trp
275 280 285

Val Ser Phe Asp Gln Asn Leu Asp Tyr Gln Ile Gly Tyr Ile Cys Ser
290 295 300

Gly Val Phe Gly Asp Asn Pro Arg Pro Lys Asp Gly Thr Gly Ser Cys
305 310 315 320

Gly Pro Val Tyr Val Asp Gly Ala Asn Gly Val Lys Gly Phe Ser Tyr
325 330 335

Arg Tyr Gly Asn Gly Val Trp Ile Gly Arg Thr Lys Ser His Ser Ser
340 345 350

Arg His Gly Phe Glu Met Ile Trp Asp Pro Asn Gly Trp Thr Glu Thr
355 360 365

Asp Ser Lys Phe Ser Val Arg Gln Asp Val Val Ala Met Thr Asp Trp
370 375 380

Ser Gly Tyr Ser Gly Ser Phe Val Gln His Pro Glu Leu Thr Gly Leu
385 390 395 400

Asp Cys Ile Arg Pro Cys Phe Trp Val Glu Leu Ile Arg Gly Arg Pro
405 410 415

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Lys Glu Lys Thr Ile Trp Thr Ser Ala Ser Ser Ile Ser Phe Cys Gly
420 425 430

Val Asn Ser Asp Thr Val Asp Trp Ser Trp Pro Asp Gly Ala Glu Leu
435 440 445

Pro Phe Thr Ile Asp Lys
450

<210> 19
<211> 566
<212> PRT
<213> Influenza A virus
<400> 19

Met Lys Ala Asn Leu Leu Val Leu Leu Cys Ala Leu Ala Ala Asp
1 5 10 15

Ala Asp Thr Ile Cys Ile Gly Tyr His Ala Asn Asn Ser Thr Asp Thr
20 25 30

Val Asp Thr Val Leu Glu Lys Asn Val Thr Val Thr His Ser Val Asn
35 40 45

Leu Leu Glu Asp Ser His Asn Gly Lys Leu Cys Arg Leu Lys Gly Ile
50 55 60

Ala Pro Leu Gln Leu Gly Lys Cys Asn Ile Ala Gly Trp Leu Leu Gly
65 70 75 80

Asn Pro Glu Cys Asp Pro Leu Leu Pro Val Arg Ser Trp Ser Tyr Ile
85 90 95

Val Glu Thr Pro Asn Ser Glu Asn Gly Ile Cys Tyr Pro Gly Asp Phe
100 105 110

Ile Asp Tyr Glu Glu Leu Arg Glu Gln Leu Ser Ser Val Ser Ser Phe
115 120 125

Glu Arg Phe Glu Ile Phe Pro Lys Glu Ser Ser Trp Pro Asn His Asn
130 135 140

Thr Thr Lys Gly Val Thr Ala Ala Cys Ser His Ala Gly Lys Ser Ser
145 150 155 160

Phe Tyr Arg Asn Leu Leu Trp Leu Thr Glu Lys Glu Gly Ser Tyr Pro
165 170 175

Lys Leu Lys Asn Ser Tyr Val Asn Lys Lys Gly Lys Glu Val Leu Val
180 185 190

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Leu Trp Gly Ile His His Pro Ser Asn Ser Lys Asp Gln Gln Asn Ile
 195 200 205
 Tyr Gln Asn Glu Asn Ala Tyr Val Ser Val Val Thr Ser Asn Tyr Asn
 210 215 220
 Arg Arg Phe Thr Pro Glu Ile Ala Glu Arg Pro Lys Val Arg Asp Gln
 225 230 235 240
 Ala Gly Arg Met Asn Tyr Tyr Trp Thr Leu Leu Lys Pro Gly Asp Thr
 245 250 255
 Ile Ile Phe Glu Ala Asn Gly Asn Leu Ile Ala Pro Arg Tyr Ala Phe
 260 265 270
 Ala Leu Ser Arg Gly Phe Gly Ser Gly Ile Ile Thr Ser Asn Ala Ser
 275 280 285
 Met His Glu Cys Asn Thr Lys Cys Gln Thr Pro Leu Gly Ala Ile Asn
 290 295 300
 Ser Ser Leu Pro Phe Gln Asn Ile His Pro Val Thr Ile Gly Glu Cys
 305 310 315 320
 Pro Lys Tyr Val Arg Ser Ala Lys Leu Arg Met Val Thr Gly Leu Arg
 325 330 335
 Asn Ile Pro Ser Ile Gln Ser Arg Gly Leu Phe Gly Ala Ile Ala Gly
 340 345 350
 Phe Ile Glu Gly Gly Trp Thr Gly Met Ile Asp Gly Trp Tyr Gly Tyr
 355 360 365
 His His Gln Asn Glu Gln Gly Ser Gly Tyr Ala Ala Asp Gln Lys Ser
 370 375 380
 Thr Gln Asn Ala Ile Asn Gly Ile Thr Asn Lys Val Asn Ser Val Ile
 385 390 395 400
 Glu Lys Met Asn Ile Gln Phe Thr Ala Val Gly Lys Glu Phe Asn Lys
 405 410 415
 Leu Glu Lys Arg Met Glu Asn Leu Asn Lys Lys Val Asp Asp Gly Phe
 420 425 430
 Leu Asp Ile Trp Thr Tyr Asn Ala Glu Leu Leu Val Leu Leu Glu Asn
 435 440 445
 Glu Arg Thr Leu Asp Phe His Asp Ser Asn Val Lys Asn Leu Tyr Glu
 450 455 460

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Lys Val Lys Ser Gln Leu Lys Asn Asn Ala Lys Glu Ile Gly Asn Gly
465 470 475 480

Cys Phe Glu Phe Tyr His Lys Cys Asp Asn Glu Cys Met Glu Ser Val
485 490 495

Arg Asn Gly Thr Tyr Asp Tyr Pro Lys Tyr Ser Glu Glu Ser Lys Leu
500 505 510

Asn Arg Glu Lys Val Asp Gly Val Lys Leu Glu Ser Met Gly Ile Tyr
515 520 525

Gln Ile Leu Ala Ile Tyr Ser Thr Val Ala Ser Ser Leu Val Leu Leu
530 535 540

Val Ser Leu Gly Ala Ile Ser Phe Trp Met Cys Ser Asn Gly Ser Leu
545 550 555 560

Gln Cys Arg Ile Cys Ile
565

<210> 20
<211> 252
<212> PRT
<213> Influenza A virus

<400> 20

Met Ser Leu Leu Thr Glu Val Glu Thr Tyr Val Leu Ser Ile Ile Pro
1 5 10 15

Ser Gly Pro Leu Lys Ala Glu Ile Ala Gln Arg Leu Glu Asp Val Phe
20 25 30

Ala Gly Lys Asn Thr Asp Leu Glu Val Leu Met Glu Trp Leu Lys Thr
35 40 45

Arg Pro Ile Leu Ser Pro Leu Thr Lys Gly Ile Leu Gly Phe Val Phe
50 55 60

Thr Leu Thr Val Pro Ser Glu Arg Gly Leu Gln Arg Arg Arg Phe Val
65 70 75 80

Gln Asn Ala Leu Asn Gly Asn Gly Asp Pro Asn Asn Met Asp Lys Ala
85 90 95

Val Lys Leu Tyr Arg Lys Leu Lys Arg Glu Ile Thr Phe His Gly Ala
100 105 110

Lys Glu Ile Ser Leu Ser Tyr Ser Ala Gly Ala Leu Ala Ser Cys Met
115 120 125

pctdk2011050460-seq1

Gly Leu Ile Tyr Asn Arg Met Gly Ala Val Thr Thr Glu Val Ala Phe
130 135 140

Gly Leu Val Cys Ala Thr Cys Glu Gln Ile Ala Asp Ser Gln His Arg
145 150 155 160

Ser His Arg Gln Met Val Thr Thr Thr Asn Pro Leu Ile Arg His Glu
165 170 175

Asn Arg Met Val Leu Ala Ser Thr Thr Ala Lys Ala Met Glu Gln Met
180 185 190

Ala Gly Ser Ser Glu Gln Ala Ala Glu Ala Met Glu Val Ala Ser Gln
195 200 205

Ala Arg Gln Met Val Gln Ala Met Arg Thr Ile Gly Thr His Pro Ser
210 215 220

Ser Ser Ala Gly Leu Lys Asn Asp Leu Leu Glu Asn Leu Gln Ala Tyr
225 230 235 240

Gln Lys Arg Met Gly Val Gln Met Gln Arg Phe Lys
245 250

<210> 21
<211> 87
<212> PRT
<213> Influenza A virus

<400> 21

Met Gly Gln Glu Gln Asp Thr Pro Trp Ile Leu Ser Thr Gly His Ile
1 5 10 15

Ser Thr Gln Lys Arg Gln Asp Gly Gln Gln Thr Pro Lys Leu Glu His
20 25 30

Arg Asn Ser Thr Arg Leu Met Gly His Cys Gln Lys Thr Met Asn Gln
35 40 45

Val Val Met Pro Lys Gln Ile Val Tyr Trp Lys Gln Trp Leu Ser Leu
50 55 60

Arg Asn Pro Ile Leu Val Phe Leu Lys Thr Arg Val Leu Lys Arg Trp
65 70 75 80

Arg Leu Phe Ser Lys His Glu
85

<210> 22
<211> 757
<212> PRT
<213> Influenza A virus

pctdk2011050460-seql

<400> 22

Met Asp Val Asn Pro Thr Leu Leu Phe Leu Lys Val Pro Ala Gln Asn
 1 5 10 15
 Ala Ile Ser Thr Thr Phe Pro Tyr Thr Gly Asp Pro Pro Tyr Ser His
 20 25 30
 Gly Thr Gly Thr Gly Tyr Thr Met Asp Thr Val Asn Arg Thr His Gln
 35 40 45
 Tyr Ser Glu Lys Ala Arg Trp Thr Thr Asn Thr Glu Thr Gly Ala Pro
 50 55 60
 Gln Leu Asn Pro Ile Asp Gly Pro Leu Pro Glu Asp Asn Glu Pro Ser
 65 70 75 80
 Gly Tyr Ala Gln Thr Asp Cys Val Leu Glu Ala Met Ala Phe Leu Glu
 85 90 95
 Glu Ser His Pro Gly Ile Phe Glu Asn Ser Cys Ile Glu Thr Met Glu
 100 105 110
 Val Val Gln Gln Thr Arg Val Asp Lys Leu Thr Gln Gly Arg Gln Thr
 115 120 125
 Tyr Asp Trp Thr Leu Asn Arg Asn Gln Pro Ala Ala Thr Ala Leu Ala
 130 135 140
 Asn Thr Ile Glu Val Phe Arg Ser Asn Gly Leu Thr Ala Asn Glu Ser
 145 150 155 160
 Gly Arg Leu Ile Asp Phe Leu Lys Asp Val Met Glu Ser Met Lys Lys
 165 170 175
 Glu Glu Met Gly Ile Thr Thr His Phe Gln Arg Lys Arg Arg Val Arg
 180 185 190
 Asp Asn Met Thr Lys Lys Met Ile Thr Gln Arg Thr Ile Gly Lys Arg
 195 200 205
 Lys Gln Arg Leu Asn Lys Arg Ser Tyr Leu Ile Arg Ala Leu Thr Leu
 210 215 220
 Asn Thr Met Thr Lys Asp Ala Glu Arg Gly Lys Leu Lys Arg Arg Ala
 225 230 235 240
 Ile Ala Thr Pro Gly Met Gln Ile Arg Gly Phe Val Tyr Phe Val Glu
 245 250 255
 Thr Leu Ala Arg Ser Ile Cys Glu Lys Leu Glu Gln Ser Gly Leu Pro
 260 265 270

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Val Gly Gly Asn Glu Lys Lys Ala Lys Leu Ala Asn Val Val Arg Lys
275 280 285
Met Met Thr Asn Ser Gln Asp Thr Glu Leu Ser Leu Thr Ile Thr Gly
290 295 300
Asp Asn Thr Lys Trp Asn Glu Asn Gln Asn Pro Arg Met Phe Leu Ala
305 310 315 320
Met Ile Thr Tyr Met Thr Arg Asn Gln Pro Glu Trp Phe Arg Asn Val
325 330 335
Leu Ser Ile Ala Pro Ile Met Phe Ser Asn Lys Met Ala Arg Leu Gly
340 345 350
Lys Gly Tyr Met Phe Glu Ser Lys Ser Met Lys Leu Arg Thr Gln Ile
355 360 365
Pro Ala Glu Met Leu Ala Ser Ile Asp Leu Lys Tyr Phe Asn Asp Ser
370 375 380
Thr Arg Lys Lys Ile Glu Lys Ile Arg Pro Leu Leu Ile Glu Gly Thr
385 390 395 400
Ala Ser Leu Ser Pro Gly Met Met Met Gly Met Phe Asn Met Leu Ser
405 410 415
Thr Val Leu Gly Val Ser Ile Leu Asn Leu Gly Gln Lys Arg Tyr Thr
420 425 430
Lys Thr Thr Tyr Trp Trp Asp Gly Leu Gln Ser Ser Asp Asp Phe Ala
435 440 445
Leu Ile Val Asn Ala Pro Asn His Glu Gly Ile Gln Ala Gly Val Asp
450 455 460
Arg Phe Tyr Arg Thr Cys Lys Leu His Gly Ile Asn Met Ser Lys Lys
465 470 475 480
Lys Ser Tyr Ile Asn Arg Thr Gly Thr Phe Glu Phe Thr Ser Phe Phe
485 490 495
Tyr Arg Tyr Gly Phe Val Ala Asn Phe Ser Met Glu Leu Pro Ser Phe
500 505 510
Gly Val Ser Gly Ser Asn Glu Ser Ala Asp Met Ser Ile Gly Val Thr
515 520 525
Val Ile Lys Asn Asn Met Ile Asn Asn Asp Leu Gly Pro Ala Thr Ala
530 535 540

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Gln Met Ala Leu Gln Leu Phe Ile Lys Asp Tyr Arg Tyr Thr Tyr Arg
545 550 555 560

Cys His Arg Gly Asp Thr Gln Ile Gln Thr Arg Arg Ser Phe Glu Ile
565 570 575

Lys Lys Leu Trp Glu Gln Thr Arg Ser Lys Ala Gly Leu Leu Val Ser
580 585 590

Asp Gly Gly Pro Asn Leu Tyr Asn Ile Arg Asn Leu His Ile Pro Glu
595 600 605

Val Cys Leu Lys Trp Glu Leu Met Asp Glu Asp Tyr Gln Gly Arg Leu
610 615 620

Cys Asn Pro Leu Asn Pro Phe Val Ser His Lys Glu Ile Glu Ser Met
625 630 635 640

Asn Asn Ala Val Met Met Pro Ala His Gly Pro Ala Lys Asn Met Glu
645 650 655

Tyr Asp Ala Val Ala Thr Thr His Ser Trp Ile Pro Lys Arg Asn Arg
660 665 670

Ser Ile Leu Asn Thr Ser Gln Arg Gly Val Leu Glu Asp Glu Gln Met
675 680 685

Tyr Gln Arg Cys Cys Asn Leu Phe Glu Lys Phe Phe Pro Ser Ser Ser
690 695 700

Tyr Arg Arg Pro Val Gly Ile Ser Ser Met Val Glu Ala Met Val Ser
705 710 715 720

Arg Ala Arg Ile Asp Ala Arg Ile Asp Phe Glu Ser Gly Arg Ile Lys
725 730 735

Lys Glu Glu Phe Thr Glu Ile Met Lys Ile Cys Ser Thr Ile Glu Glu
740 745 750

Leu Arg Arg Gln Lys
755

<210> 23
<211> 498
<212> PRT
<213> Influenza A virus

<400> 23

Met Ala Ser Gln Gly Thr Lys Arg Ser Tyr Glu Gln Met Glu Thr Asp
1 5 10 15

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Gly Glu Arg Gln Asn Ala Thr Glu Ile Arg Ala Ser Val Gly Lys Met
20 25 30
Ile Gly Gly Ile Gly Arg Phe Tyr Ile Gln Met Cys Thr Glu Leu Lys
35 40 45
Leu Ser Asp Tyr Glu Gly Arg Leu Ile Gln Asn Ser Leu Thr Ile Glu
50 55 60
Arg Met Val Leu Ser Ala Phe Asp Glu Arg Arg Asn Lys Tyr Leu Glu
65 70 75 80
Glu His Pro Ser Ala Gly Lys Asp Pro Lys Lys Thr Gly Gly Pro Ile
85 90 95
Tyr Arg Arg Val Asn Gly Lys Trp Met Arg Glu Leu Ile Leu Tyr Asp
100 105 110
Lys Glu Glu Ile Arg Arg Ile Trp Arg Gln Ala Asn Asn Gly Asp Asp
115 120 125
Ala Thr Ala Gly Leu Thr His Met Met Ile Trp His Ser Asn Leu Asn
130 135 140
Asp Ala Thr Tyr Gln Arg Thr Arg Ala Leu Val Arg Thr Gly Met Asp
145 150 155 160
Pro Arg Met Cys Ser Leu Met Gln Gly Ser Thr Leu Pro Arg Arg Ser
165 170 175
Gly Ala Ala Gly Ala Ala Val Lys Gly Val Gly Thr Met Val Met Glu
180 185 190
Leu Val Arg Met Ile Lys Arg Gly Ile Asn Asp Arg Asn Phe Trp Arg
195 200 205
Gly Glu Asn Gly Arg Lys Thr Arg Ile Ala Tyr Glu Arg Met Cys Asn
210 215 220
Ile Leu Lys Gly Lys Phe Gln Thr Ala Ala Gln Lys Ala Met Met Asp
225 230 235 240
Gln Val Arg Glu Ser Arg Asp Pro Gly Asn Ala Glu Phe Glu Asp Leu
245 250 255
Thr Phe Leu Ala Arg Ser Ala Leu Ile Leu Arg Gly Ser Val Ala His
260 265 270
Lys Ser Cys Leu Pro Ala Cys Val Tyr Gly Pro Ala Val Ala Ser Gly
275 280 285

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Tyr Asp Phe Glu Arg Glu Gly Tyr Ser Leu Val Gly Ile Asp Pro Phe
290 295 300

Arg Leu Leu Gln Asn Ser Gln Val Tyr Ser Leu Ile Arg Pro Asn Glu
305 310 315 320

Asn Pro Ala His Lys Ser Gln Leu Val Trp Met Ala Cys His Ser Ala
325 330 335

Ala Phe Glu Asp Leu Arg Val Leu Ser Phe Ile Lys Gly Thr Lys Val
340 345 350

Val Pro Arg Gly Lys Leu Ser Thr Arg Gly Val Gln Ile Ala Ser Asn
355 360 365

Glu Asn Met Glu Thr Met Glu Ser Ser Thr Leu Glu Leu Arg Ser Arg
370 375 380

Tyr Trp Ala Ile Arg Thr Arg Ser Gly Gly Asn Thr Asn Gln Gln Arg
385 390 395 400

Ala Ser Ala Gly Gln Ile Ser Ile Gln Pro Thr Phe Ser Val Gln Arg
405 410 415

Asn Leu Pro Phe Asp Arg Thr Thr Val Met Ala Ala Phe Thr Gly Asn
420 425 430

Thr Glu Gly Arg Thr Ser Asp Met Arg Thr Glu Ile Ile Arg Met Met
435 440 445

Glu Ser Ala Arg Pro Glu Asp Val Ser Phe Gln Gly Arg Gly Val Phe
450 455 460

Glu Leu Ser Asp Glu Lys Ala Ala Ser Pro Ile Val Pro Ser Phe Asp
465 470 475 480

Met Ser Asn Glu Gly Ser Tyr Phe Phe Gly Asp Asn Ala Glu Glu Tyr
485 490 495

Asp Asn

<210> 24
<211> 759
<212> PRT
<213> Influenza A virus

<400> 24

Met Glu Arg Ile Lys Glu Leu Arg Asn Leu Met Ser Gln Ser Arg Thr
1 5 10 15

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Arg Glu Ile Leu Thr Lys Thr Thr Val Asp His Met Ala Ile Ile Lys
20 25 30

Lys Tyr Thr Ser Gly Arg Gln Glu Lys Asn Pro Ser Leu Arg Met Lys
35 40 45

Trp Met Met Ala Met Lys Tyr Pro Ile Thr Ala Asp Lys Arg Ile Thr
50 55 60

Glu Met Val Pro Glu Arg Asn Glu Gln Gly Gln Thr Leu Trp Ser Lys
65 70 75 80

Met Ser Asp Ala Gly Ser Asp Arg Val Met Val Ser Pro Leu Ala Val
85 90 95

Thr Trp Trp Asn Arg Asn Gly Pro Met Thr Ser Thr Val His Tyr Pro
100 105 110

Lys Ile Tyr Lys Thr Tyr Phe Glu Lys Val Glu Arg Leu Lys His Gly
115 120 125

Thr Phe Gly Pro Val His Phe Arg Asn Gln Val Lys Ile Arg Arg Arg
130 135 140

Val Asp Ile Asn Pro Gly His Ala Asp Leu Ser Ala Lys Glu Ala Gln
145 150 155 160

Asp Val Ile Met Glu Val Val Phe Pro Asn Glu Val Gly Ala Arg Ile
165 170 175

Leu Thr Ser Glu Ser Gln Leu Thr Ile Thr Lys Glu Lys Lys Glu Glu
180 185 190

Leu Gln Asp Cys Lys Ile Ser Pro Leu Met Val Ala Tyr Met Leu Glu
195 200 205

Arg Glu Leu Val Arg Lys Thr Arg Phe Leu Pro Val Ala Gly Gly Thr
210 215 220

Ser Ser Val Tyr Ile Glu Val Leu His Leu Thr Gln Gly Thr Cys Trp
225 230 235 240

Glu Gln Met Tyr Thr Pro Gly Gly Glu Val Arg Asn Asp Asp Val Asp
245 250 255

Gln Ser Leu Ile Ile Ala Ala Arg Asn Ile Val Arg Arg Ala Ala Val
260 265 270

Ser Ala Asp Pro Leu Ala Ser Leu Leu Glu Met Cys His Ser Thr Gln
275 280 285

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Ile	Gly	Gly	Thr	Arg	Met	Val	Asp	Ile	Leu	Arg	Gln	Asn	Pro	Thr	Glu
	290					295					300				
Glu	Gln	Ala	Val	Asp	Ile	Cys	Lys	Ala	Ala	Met	Gly	Leu	Arg	Ile	Ser
305					310					315					320
Ser	Ser	Phe	Ser	Phe	Gly	Gly	Phe	Thr	Phe	Lys	Arg	Thr	Ser	Gly	Ser
				325					330					335	
Ser	Ile	Lys	Arg	Glu	Glu	Glu	Val	Leu	Thr	Gly	Asn	Leu	Gln	Thr	Leu
			340					345					350		
Lys	Ile	Arg	Val	His	Glu	Gly	Tyr	Glu	Glu	Phe	Thr	Met	Val	Gly	Lys
		355					360					365			
Arg	Ala	Thr	Ala	Ile	Leu	Arg	Lys	Ala	Thr	Arg	Arg	Leu	Val	Gln	Leu
	370					375					380				
Ile	Val	Ser	Gly	Arg	Asp	Glu	Gln	Ser	Ile	Ala	Glu	Ala	Ile	Ile	Val
385					390					395					400
Ala	Met	Val	Phe	Ser	Gln	Glu	Asp	Cys	Met	Ile	Lys	Ala	Val	Arg	Gly
				405					410					415	
Asp	Leu	Asn	Phe	Val	Asn	Arg	Ala	Asn	Gln	Arg	Leu	Asn	Pro	Met	His
			420					425					430		
Gln	Leu	Leu	Arg	His	Phe	Gln	Lys	Asp	Ala	Lys	Val	Leu	Phe	Gln	Asn
		435					440					445			
Trp	Gly	Ile	Glu	His	Ile	Asp	Asn	Val	Met	Gly	Met	Ile	Gly	Val	Leu
	450					455					460				
Pro	Asp	Met	Thr	Pro	Ser	Thr	Glu	Met	Ser	Met	Arg	Gly	Ile	Arg	Val
465					470					475					480
Ser	Lys	Met	Gly	Val	Asp	Glu	Tyr	Ser	Ser	Thr	Glu	Arg	Val	Val	Val
				485					490					495	
Ser	Ile	Asp	Arg	Phe	Leu	Arg	Val	Arg	Asp	Gln	Arg	Gly	Asn	Val	Leu
			500					505					510		
Leu	Ser	Pro	Glu	Glu	Val	Ser	Glu	Thr	Gln	Gly	Thr	Glu	Lys	Leu	Thr
		515					520					525			
Ile	Thr	Tyr	Ser	Ser	Ser	Met	Met	Trp	Glu	Ile	Asn	Gly	Pro	Glu	Ser
	530					535					540				
Val	Leu	Val	Asn	Thr	Tyr	Gln	Trp	Ile	Ile	Arg	Asn	Trp	Glu	Thr	Val
545					550					555					560

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Lys Ile Gln Trp Ser Gln Asn Pro Thr Met Leu Tyr Asn Lys Met Glu
565 570 575

Phe Glu Pro Phe Gln Ser Leu Val Pro Lys Ala Ile Arg Gly Gln Tyr
580 585 590

Ser Gly Phe Val Arg Thr Leu Phe Gln Gln Met Arg Asp Val Leu Gly
595 600 605

Thr Phe Asp Thr Thr Gln Ile Ile Lys Leu Leu Pro Phe Ala Ala Ala
610 615 620

Pro Pro Lys Gln Ser Arg Met Gln Phe Ser Ser Leu Thr Val Asn Val
625 630 635 640

Arg Gly Ser Gly Met Arg Ile Leu Val Arg Gly Asn Ser Pro Val Phe
645 650 655

Asn Tyr Asn Lys Thr Thr Lys Arg Leu Thr Ile Leu Gly Lys Asp Ala
660 665 670

Gly Thr Leu Thr Glu Asp Pro Asp Glu Gly Thr Ser Gly Val Glu Ser
675 680 685

Ala Val Leu Arg Gly Phe Leu Ile Leu Gly Lys Glu Asp Arg Arg Tyr
690 695 700

Gly Pro Ala Leu Ser Ile Asn Glu Leu Ser Thr Leu Ala Lys Gly Glu
705 710 715 720

Lys Ala Asn Val Leu Ile Gly Gln Gly Asp Val Val Leu Val Met Lys
725 730 735

Arg Lys Arg Asp Ser Ser Ile Leu Thr Asp Ser Gln Thr Ala Thr Lys
740 745 750

Arg Ile Arg Met Ala Ile Asn
755

<210> 25
<211> 562
<212> PRT
<213> Influenza A virus

<400> 25

Met Ala Ile Ile Tyr Leu Ile Leu Leu Phe Thr Ala Val Arg Gly Asp
1 5 10 15

Gln Ile Cys Ile Gly Tyr His Ala Asn Asn Ser Thr Glu Lys Val Asp
20 25 30

Thr Ile Leu Glu Arg Asn Val Thr Val Thr His Ala Lys Asp Ile Leu
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35

40

45

Glu Lys Thr His Asn Gly Lys Leu Cys Lys Leu Asn Gly Ile Pro Pro
 50 55 60
 Leu Glu Leu Gly Asp Cys Ser Ile Ala Gly Trp Leu Leu Gly Asn Pro
 65 70 75 80
 Glu Cys Asp Arg Leu Leu Ser Val Pro Glu Trp Ser Tyr Ile Met Glu
 85 90 95
 Lys Glu Asn Pro Arg Tyr Ser Leu Cys Tyr Pro Gly Ser Phe Asn Asp
 100 105 110
 Tyr Glu Glu Leu Lys His Leu Leu Ser Ser Val Lys His Phe Glu Lys
 115 120 125
 Val Lys Ile Leu Pro Lys Asp Arg Trp Thr Gln His Thr Thr Thr Gly
 130 135 140
 Gly Ser Trp Ala Cys Ala Val Ser Gly Lys Pro Ser Phe Phe Arg Asn
 145 150 155 160
 Met Val Trp Leu Thr Arg Lys Gly Ser Asn Tyr Pro Val Ala Lys Gly
 165 170 175
 Ser Tyr Asn Asn Thr Ser Gly Glu Gln Met Leu Ile Ile Trp Gly Val
 180 185 190
 His His Pro Asn Asp Glu Ala Glu Gln Arg Ala Leu Tyr Gln Asn Val
 195 200 205
 Gly Thr Tyr Val Ser Val Ala Thr Ser Thr Leu Tyr Lys Arg Ser Ile
 210 215 220
 Pro Glu Ile Ala Ala Arg Pro Lys Val Asn Gly Leu Gly Arg Arg Met
 225 230 235 240
 Glu Phe Ser Trp Thr Leu Leu Asp Met Trp Asp Thr Ile Asn Phe Glu
 245 250 255
 Ser Thr Gly Asn Leu Val Ala Pro Glu Tyr Gly Phe Lys Ile Ser Lys
 260 265 270
 Arg Gly Ser Ser Gly Ile Met Lys Thr Glu Gly Thr Leu Glu Asn Cys
 275 280 285
 Glu Thr Lys Cys Gln Thr Pro Leu Gly Ala Ile Asn Thr Thr Leu Pro
 290 295 300
 Phe His Asn Val His Pro Leu Thr Ile Gly Glu Cys Pro Lys Tyr Val

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305          310          315          320
Lys Ser Glu Lys Leu Val Leu Ala Thr Gly Leu Arg Asn Val Pro Gln
          325          330          335
Ile Glu Ser Arg Gly Leu Phe Gly Ala Ile Ala Gly Phe Ile Glu Gly
          340          345          350
Gly Trp Gln Gly Met Val Asp Gly Trp Tyr Gly Tyr His His Ser Asn
          355          360          365
Asp Gln Gly Ser Gly Tyr Ala Ala Asp Lys Glu Ser Thr Gln Lys Ala
          370          375          380
Phe Asn Gly Ile Thr Asn Lys Val Asn Ser Val Ile Glu Lys Met Asn
          385          390          395
Thr Gln Phe Glu Ala Val Gly Lys Glu Phe Ser Asn Leu Glu Lys Arg
          405          410          415
Leu Glu Asn Leu Asn Lys Lys Met Glu Asp Gly Phe Leu Asp Val Trp
          420          425          430
Thr Tyr Asn Ala Glu Leu Leu Val Leu Met Glu Asn Glu Arg Thr Leu
          435          440          445
Asp Phe His Asp Ser Asn Val Lys Asn Leu Tyr Asp Lys Val Arg Met
          450          455          460
Gln Leu Arg Asp Asn Val Lys Glu Leu Gly Asn Gly Cys Phe Glu Phe
          465          470          475
Tyr His Lys Cys Asp Asn Glu Cys Met Asp Ser Val Lys Asn Gly Thr
          485          490          495
Tyr Asp Tyr Pro Lys Tyr Glu Glu Glu Ser Lys Leu Asn Arg Asn Glu
          500          505          510
Ile Lys Gly Val Lys Leu Ser Ser Met Gly Val Tyr Gln Ile Leu Ala
          515          520          525
Ile Tyr Ala Thr Val Ala Gly Ser Leu Ser Leu Ala Ile Met Met Ala
          530          535          540
Gly Ile Ser Phe Trp Met Cys Ser Asn Gly Ser Leu Gln Cys Arg Ile
          545          550          555          560

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Cys Ile

<210> 26

pctdk2011050460-seq1

<211> 252
 <212> PRT
 <213> Influenza A virus

<400> 26

Met Ser Leu Leu Thr Glu Val Glu Thr Tyr Val Leu Ser Ile Val Pro
 1 5 10 15

Ser Gly Pro Leu Lys Ala Glu Ile Ala Gln Arg Leu Glu Asp Val Phe
 20 25 30

Ala Gly Lys Asn Thr Asp Leu Glu Ala Leu Met Glu Trp Leu Lys Thr
 35 40 45

Arg Pro Ile Leu Ser Pro Leu Thr Lys Gly Ile Leu Gly Phe Val Phe
 50 55 60

Thr Leu Thr Val Pro Ser Glu Arg Gly Leu Gln Arg Arg Arg Phe Val
 65 70 75 80

Gln Asn Ala Leu Asn Gly Asn Gly Asp Pro Asn Asn Met Asp Arg Ala
 85 90 95

Val Lys Leu Tyr Arg Lys Leu Lys Arg Glu Ile Thr Phe His Gly Ala
 100 105 110

Lys Glu Val Ala Leu Ser Tyr Ser Ala Gly Ala Leu Ala Ser Cys Met
 115 120 125

Gly Leu Ile Tyr Asn Arg Met Gly Ala Val Thr Thr Glu Val Ala Phe
 130 135 140

Ala Val Val Cys Ala Thr Cys Glu Gln Ile Ala Asp Ser Gln His Arg
 145 150 155 160

Ser His Arg Gln Met Val Thr Thr Thr Asn Pro Leu Ile Arg His Glu
 165 170 175

Asn Arg Met Val Leu Ala Ser Thr Thr Ala Lys Ala Met Glu Gln Met
 180 185 190

Ala Gly Ser Ser Glu Gln Ala Ala Glu Ala Met Glu Val Ala Ser Gln
 195 200 205

Ala Arg Gln Met Val Gln Ala Met Arg Ala Ile Gly Thr Pro Pro Ser
 210 215 220

Ser Ser Ala Gly Leu Lys Asp Asp Leu Leu Glu Asn Leu Gln Ala Tyr
 225 230 235 240

Gln Lys Arg Met Gly Val Gln Met Gln Arg Phe Lys
 245 250

pctdk2011050460-seq1

<210> 27
 <211> 97
 <212> PRT
 <213> Influenza A virus

<400> 27

Met Ser Leu Leu Thr Glu Val Glu Thr Pro Ile Arg Asn Glu Trp Gly
 1 5 10 15

Cys Arg Cys Asn Asp Ser Ser Asp Pro Leu Val Val Ala Ala Ser Ile
 20 25 30

Ile Gly Ile Leu His Phe Ile Leu Trp Ile Leu Asp Arg Leu Phe Phe
 35 40 45

Lys Cys Ile Tyr Arg Phe Phe Lys His Gly Leu Lys Arg Gly Pro Ser
 50 55 60

Thr Glu Gly Val Pro Glu Ser Met Arg Glu Glu Tyr Arg Lys Glu Gln
 65 70 75 80

Gln Ser Ala Val Asp Ala Asp Asp Ser His Phe Val Ser Ile Glu Leu
 85 90 95

Glu

<210> 28
 <211> 716
 <212> PRT
 <213> Influenza A virus

<400> 28

Met Glu Asp Phe Val Arg Gln Cys Phe Asn Pro Met Ile Val Glu Leu
 1 5 10 15

Ala Glu Lys Ala Met Lys Glu Tyr Gly Glu Asp Leu Lys Ile Glu Thr
 20 25 30

Asn Lys Phe Ala Ala Ile Cys Thr His Leu Glu Val Cys Phe Met Tyr
 35 40 45

Ser Asp Phe His Phe Ile Asn Glu Gln Gly Glu Ser Ile Met Val Glu
 50 55 60

Leu Asp Asp Pro Asn Ala Leu Leu Lys His Arg Phe Glu Ile Ile Glu
 65 70 75 80

Gly Arg Asp Arg Thr Met Ala Trp Thr Val Val Asn Ser Ile Cys Asn
 85 90 95

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Thr	Thr	Gly	Ala	Glu	Lys	Pro	Lys	Phe	Leu	Pro	Asp	Leu	Tyr	Asp	Tyr
			100					105					110		
Lys	Glu	Asn	Arg	Phe	Ile	Glu	Ile	Gly	Val	Thr	Arg	Arg	Glu	Val	His
		115					120					125			
Ile	Tyr	Tyr	Leu	Glu	Lys	Ala	Asn	Lys	Ile	Lys	Ser	Glu	Asn	Thr	His
	130					135					140				
Ile	His	Ile	Phe	Ser	Phe	Thr	Gly	Glu	Glu	Met	Ala	Thr	Lys	Ala	Asp
145					150					155					160
Tyr	Thr	Leu	Asp	Glu	Glu	Ser	Arg	Ala	Arg	Ile	Lys	Thr	Arg	Leu	Phe
				165					170					175	
Thr	Ile	Arg	Gln	Glu	Met	Ala	Asn	Arg	Gly	Leu	Trp	Asp	Ser	Phe	Arg
			180					185					190		
Gln	Ser	Glu	Arg	Gly	Glu	Glu	Thr	Ile	Glu	Glu	Arg	Phe	Glu	Ile	Thr
		195					200					205			
Gly	Thr	Met	Arg	Arg	Leu	Ala	Asp	Gln	Ser	Leu	Pro	Pro	Asn	Phe	Ser
	210					215					220				
Cys	Leu	Glu	Asn	Phe	Arg	Ala	Tyr	Val	Asp	Gly	Phe	Glu	Pro	Asn	Gly
225					230					235					240
Tyr	Ile	Glu	Gly	Lys	Leu	Ser	Gln	Met	Ser	Lys	Glu	Val	Asn	Ala	Lys
				245					250					255	
Ile	Glu	Pro	Phe	Leu	Lys	Thr	Thr	Pro	Arg	Pro	Ile	Arg	Leu	Pro	Asp
			260					265					270		
Gly	Pro	Pro	Cys	Phe	Gln	Arg	Ser	Lys	Phe	Leu	Leu	Met	Asp	Ala	Leu
		275					280					285			
Lys	Leu	Ser	Ile	Glu	Asp	Pro	Ser	His	Glu	Gly	Glu	Gly	Ile	Pro	Leu
	290					295					300				
Tyr	Asp	Ala	Ile	Lys	Cys	Met	Arg	Thr	Phe	Phe	Gly	Trp	Lys	Glu	Pro
305					310					315					320
Tyr	Ile	Val	Lys	Pro	His	Glu	Lys	Gly	Ile	Asn	Pro	Asn	Tyr	Leu	Leu
				325					330					335	
Ser	Trp	Lys	Gln	Val	Leu	Ala	Glu	Leu	Gln	Asp	Ile	Glu	Asn	Glu	Glu
			340					345					350		
Lys	Ile	Pro	Arg	Thr	Lys	Asn	Met	Lys	Lys	Thr	Ser	Gln	Leu	Lys	Trp
		355					360					365			

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Ala Leu Gly Glu Asn Met Ala Pro Glu Lys Val Asp Phe Asp Asn Cys
370 375 380
Arg Asp Ile Ser Asp Leu Lys Gln Tyr Asp Ser Asp Glu Pro Glu Leu
385 390 395 400
Arg Ser Leu Ser Ser Trp Ile Gln Asn Glu Phe Asn Lys Ala Cys Glu
405 410 415
Leu Thr Asp Ser Ile Trp Ile Glu Leu Asp Glu Ile Gly Glu Asp Val
420 425 430
Ala Pro Ile Glu His Ile Ala Ser Met Arg Arg Asn Tyr Phe Thr Ala
435 440 445
Glu Val Ser His Cys Arg Ala Thr Glu Tyr Ile Met Lys Gly Val Tyr
450 455 460
Ile Asn Thr Ala Leu Leu Asn Ala Ser Cys Ala Ala Met Asp Asp Phe
465 470 475 480
Gln Leu Ile Pro Met Ile Ser Lys Cys Arg Thr Lys Glu Gly Arg Arg
485 490 495
Lys Thr Asn Leu Tyr Gly Phe Ile Ile Lys Gly Arg Ser His Leu Arg
500 505 510
Asn Asp Thr Asp Val Val Asn Phe Val Ser Met Glu Phe Ser Leu Thr
515 520 525
Asp Pro Arg Leu Glu Pro His Lys Trp Glu Lys Tyr Cys Val Leu Glu
530 535 540
Ile Gly Asp Met Leu Leu Arg Ser Ala Ile Gly Gln Met Ser Arg Pro
545 550 555 560
Met Phe Leu Tyr Val Arg Thr Asn Gly Thr Ser Lys Ile Lys Met Lys
565 570 575
Trp Gly Met Glu Met Arg Pro Cys Leu Leu Gln Ser Leu Gln Gln Ile
580 585 590
Glu Ser Met Val Glu Ala Glu Ser Ser Val Lys Glu Lys Asp Met Thr
595 600 605
Lys Glu Phe Phe Glu Asn Lys Ser Glu Thr Trp Pro Ile Gly Glu Ser
610 615 620
Pro Lys Gly Val Glu Glu Gly Ser Ile Gly Lys Val Cys Arg Thr Leu
625 630 635 640

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Leu Ala Lys Ser Val Phe Asn Ser Leu Tyr Ala Ser Pro Gln Leu Glu
645 650 655

Gly Phe Ser Ala Glu Ser Arg Lys Leu Leu Leu Val Val Gln Ala Leu
660 665 670

Arg Asp Asn Leu Glu Pro Gly Thr Phe Asp Leu Gly Gly Leu Tyr Glu
675 680 685

Ala Ile Glu Glu Cys Leu Ile Asn Asp Pro Trp Val Leu Leu Asn Ala
690 695 700

Ser Trp Phe Asn Ser Phe Leu Thr His Ala Leu Arg
705 710 715

<210> 29
<211> 90
<212> PRT
<213> Influenza A virus

<400> 29

Met Gly Gln Glu Gln Asp Thr Pro Trp Thr Gln Ser Thr Glu His Ile
1 5 10 15

Asn Ile Gln Lys Arg Gly Ser Gly Gln Gln Thr Arg Lys Leu Glu Arg
20 25 30

Pro Asn Leu Thr Gln Leu Met Asp His Tyr Leu Arg Thr Met Asn Gln
35 40 45

Val Asp Met His Lys Gln Thr Ala Ser Trp Lys Gln Trp Leu Ser Leu
50 55 60

Arg Asn His Thr Gln Glu Ser Leu Lys Ile Arg Val Leu Lys Arg Trp
65 70 75 80

Lys Leu Phe Asn Lys Gln Glu Trp Thr Asn
85 90

<210> 30
<211> 757
<212> PRT
<213> Influenza A virus

<400> 30

Met Asp Val Asn Pro Thr Leu Leu Phe Leu Lys Val Pro Ala Gln Asn
1 5 10 15

Ala Ile Ser Thr Thr Phe Pro Tyr Thr Gly Asp Pro Pro Tyr Ser His
20 25 30

Gly Thr Gly Thr Gly Tyr Thr Met Asp Thr Val Asn Arg Thr His Gln
35 40 45

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Tyr Ser Glu Lys Gly Lys Trp Thr Thr Asn Thr Glu Thr Gly Ala Pro
50 55 60
Gln Leu Asn Pro Ile Asp Gly Pro Leu Pro Glu Asp Asn Glu Pro Ser
65 70 75 80
Gly Tyr Ala Gln Thr Asp Cys Val Leu Glu Ala Met Ala Phe Leu Glu
85 90 95
Glu Ser His Pro Gly Ile Phe Glu Asn Ser Cys Leu Glu Thr Met Glu
100 105 110
Val Ile Gln Gln Thr Arg Val Asp Lys Leu Thr Gln Gly Arg Gln Thr
115 120 125
Tyr Asp Trp Thr Leu Asn Arg Asn Gln Pro Ala Ala Thr Ala Leu Ala
130 135 140
Asn Thr Ile Glu Val Phe Arg Ser Asn Gly Leu Thr Ala Asn Glu Ser
145 150 155 160
Gly Arg Leu Ile Asp Phe Leu Lys Asp Val Ile Glu Ser Met Asp Lys
165 170 175
Glu Glu Met Glu Ile Thr Thr His Phe Gln Arg Lys Arg Arg Val Arg
180 185 190
Asp Asn Met Thr Lys Lys Met Val Thr Gln Arg Thr Ile Gly Lys Lys
195 200 205
Lys Gln Arg Leu Asn Lys Arg Ser Tyr Leu Ile Arg Ala Leu Thr Leu
210 215 220
Asn Thr Met Thr Lys Asp Ala Glu Arg Gly Lys Leu Lys Arg Arg Ala
225 230 235 240
Ile Ala Thr Pro Gly Met Gln Ile Arg Gly Phe Val His Phe Val Glu
245 250 255
Thr Leu Ala Arg Asn Ile Cys Glu Lys Leu Glu Gln Ser Gly Leu Pro
260 265 270
Val Gly Gly Asn Glu Lys Lys Ala Lys Leu Ala Asn Val Val Arg Lys
275 280 285
Met Met Thr Asn Ser Gln Asp Thr Glu Leu Ser Phe Thr Ile Thr Gly
290 295 300
Asp Asn Thr Lys Trp Asn Glu Asn Gln Asn Pro Arg Val Phe Leu Ala
305 310 315 320

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Met Ile Thr Tyr Ile Thr Arg Asn Gln Pro Glu Trp Phe Arg Asn Val
          325          330          335

Leu Ser Ile Ala Pro Ile Met Phe Ser Asn Lys Met Ala Arg Leu Gly
          340          345          350

Lys Gly Tyr Met Phe Glu Ser Lys Ser Met Lys Leu Arg Thr Gln Ile
          355          360          365

Pro Ala Glu Met Leu Ala Ser Ile Asp Leu Lys Tyr Phe Asn Glu Ser
          370          375          380

Thr Arg Lys Lys Ile Glu Lys Ile Arg Pro Leu Leu Ile Asp Gly Thr
385          390          395          400

Val Ser Leu Ser Pro Gly Met Met Met Gly Met Phe Asn Met Leu Ser
          405          410          415

Thr Val Leu Gly Val Ser Ile Leu Asn Leu Gly Gln Lys Lys Tyr Thr
          420          425          430

Lys Thr Thr Tyr Trp Trp Asp Gly Leu Gln Ser Ser Asp Asp Phe Ala
          435          440          445

Leu Ile Val Asn Ala Pro Asn His Glu Gly Ile Gln Ala Gly Val Asn
450          455          460

Arg Phe Tyr Arg Thr Cys Lys Leu Val Gly Ile Asn Met Ser Lys Lys
465          470          475          480

Lys Ser Tyr Ile Asn Arg Thr Gly Thr Phe Glu Phe Thr Ser Phe Phe
          485          490          495

Tyr Arg Tyr Gly Phe Val Ala Asn Phe Ser Met Glu Leu Pro Ser Phe
          500          505          510

Gly Val Ser Gly Ile Asn Glu Ser Ala Asp Met Ser Ile Gly Val Thr
          515          520          525

Val Ile Lys Asn Asn Met Ile Asn Asn Asp Leu Gly Pro Ala Thr Ala
530          535          540

Gln Met Ala Leu Gln Leu Phe Ile Lys Asp Tyr Arg Tyr Thr Tyr Arg
545          550          555          560

Cys His Arg Gly Asp Thr Gln Ile Gln Thr Arg Arg Ser Phe Glu Leu
          565          570          575

Lys Lys Leu Trp Glu Gln Thr Arg Ser Lys Ala Gly Leu Leu Val Ser
          580          585          590

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Asp Gly Gly Ser Asn Leu Tyr Asn Ile Arg Asn Leu His Ile Pro Glu
595 600 605

Val Cys Leu Lys Trp Glu Leu Met Asp Glu Asp Tyr Gln Gly Arg Leu
610 615 620

Cys Asn Pro Leu Asn Pro Phe Val Ser His Lys Glu Ile Glu Ser Val
625 630 635 640

Asn Asn Ala Val Val Met Pro Ala His Gly Pro Ala Lys Ser Met Glu
645 650 655

Tyr Asp Ala Val Ala Thr Thr His Ser Trp Thr Pro Lys Arg Asn Arg
660 665 670

Ser Ile Leu Asn Thr Ser Gln Arg Gly Ile Leu Glu Asp Glu Gln Met
675 680 685

Tyr Gln Lys Cys Cys Asn Leu Phe Glu Lys Phe Phe Pro Ser Ser Ser
690 695 700

Tyr Arg Arg Pro Val Gly Ile Ser Ser Met Val Glu Ala Met Val Ser
705 710 715 720

Arg Ala Arg Ile Asp Ala Arg Ile Asp Phe Glu Ser Gly Arg Ile Lys
725 730 735

Lys Glu Glu Phe Ala Glu Ile Met Lys Ile Cys Ser Thr Ile Glu Glu
740 745 750

Leu Arg Arg Gln Lys
755

<210> 31
<211> 121
<212> PRT
<213> Influenza A virus

<400> 31

Met Asp Ser Asn Thr Val Ser Ser Phe Gln Asp Ile Leu Leu Arg Met
1 5 10 15

Ser Lys Met Gln Leu Gly Ser Ser Ser Glu Asp Leu Asn Gly Met Ile
20 25 30

Thr Gln Phe Glu Ser Leu Lys Leu Tyr Arg Asp Ser Leu Gly Glu Ala
35 40 45

Val Met Arg Met Gly Asp Leu His Ser Leu Gln Asn Arg Asn Gly Lys
50 55 60

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Trp Arg Glu Gln Leu Gly Gln Lys Phe Glu Glu Ile Arg Trp Leu Ile
65 70 75 80

Glu Glu Val Arg His Arg Leu Lys Ile Thr Glu Asn Ser Phe Glu Gln
85 90 95

Ile Thr Phe Met Gln Ala Leu Gln Leu Leu Phe Glu Val Glu Gln Glu
100 105 110

Ile Arg Thr Phe Ser Phe Gln Leu Ile
115 120

<210> 32
<211> 237
<212> PRT
<213> Influenza A virus

<400> 32

Met Asp Ser Asn Thr Val Ser Ser Phe Gln Val Asp Cys Phe Leu Trp
1 5 10 15

His Val Arg Lys Gln Val Val Asp Gln Glu Leu Gly Asp Ala Pro Phe
20 25 30

Leu Asp Arg Leu Arg Arg Asp Gln Lys Ser Leu Arg Gly Arg Gly Ser
35 40 45

Thr Leu Asp Leu Asp Ile Glu Ala Ala Thr Arg Val Gly Lys Gln Ile
50 55 60

Val Glu Arg Ile Leu Lys Glu Glu Ser Asp Glu Ala Leu Lys Met Thr
65 70 75 80

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

Glu Leu Ser Arg Asp Trp Phe Met Leu Met Pro Lys Gln Lys Val Glu
100 105 110

Gly Pro Leu Cys Ile Arg Ile Asp Gln Ala Ile Met Asp Lys Asn Ile
115 120 125

Met Leu Lys Ala Asn Phe Ser Val Ile Phe Asp Arg Leu Glu Thr Leu
130 135 140

Ile Leu Leu Arg Ala Phe Thr Glu Glu Gly Ala Ile Val Gly Glu Ile
145 150 155 160

Ser Pro Leu Pro Ser Leu Pro Gly His Thr Ile Glu Asp Val Lys Asn
165 170 175

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Ala Ile Gly Val Leu Ile Gly Gly Leu Glu Trp Asn Asp Asn Thr Val
180 185 190

Arg Val Ser Lys Thr Leu Gln Arg Phe Ala Trp Arg Ser Ser Asn Glu
195 200 205

Asn Gly Arg Pro Pro Leu Thr Pro Lys Gln Lys Arg Lys Met Ala Arg
210 215 220

Thr Ile Arg Ser Lys Val Arg Arg Asp Lys Met Ala Asp
225 230 235

<210> 33

<211> 498

<212> PRT

<213> Influenza A virus

<400> 33

Met Ala Ser Gln Gly Thr Lys Arg Ser Tyr Glu Gln Met Glu Thr Asp
1 5 10 15

Gly Glu Arg Gln Asn Ala Thr Glu Ile Arg Ala Ser Val Gly Lys Met
20 25 30

Ile Asp Gly Ile Gly Arg Phe Tyr Ile Gln Met Cys Thr Glu Leu Lys
35 40 45

Leu Ser Asp Tyr Glu Gly Arg Leu Ile Gln Asn Ser Leu Thr Ile Glu
50 55 60

Arg Met Val Leu Ser Ala Phe Asp Glu Arg Arg Asn Lys Tyr Leu Glu
65 70 75 80

Glu His Pro Ser Ala Gly Lys Asp Pro Lys Lys Thr Gly Gly Pro Ile
85 90 95

Tyr Lys Arg Val Asp Gly Lys Trp Met Arg Glu Leu Val Leu Tyr Asp
100 105 110

Lys Glu Glu Ile Arg Arg Ile Trp Arg Gln Ala Asn Asn Gly Asp Asp
115 120 125

Ala Thr Ala Gly Leu Thr His Met Met Ile Trp His Ser Asn Leu Asn
130 135 140

Asp Thr Thr Tyr Gln Arg Thr Arg Ala Leu Val Arg Thr Gly Met Asp
145 150 155 160

Pro Arg Met Cys Ser Leu Met Gln Gly Ser Thr Leu Pro Arg Arg Ser
165 170 175

Gly Ala Ala Gly Ala Ala Val Lys Gly Val Gly Thr Met Val Met Glu
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180

185

190

Leu Ile Arg Met Ile Lys Arg Gly Ile Asn Asp Arg Asn Phe Trp Arg
 195 200 205

Gly Glu Asn Gly Arg Lys Thr Arg Ser Ala Tyr Glu Arg Met Cys Asn
 210 215 220

Ile Leu Lys Gly Lys Phe Gln Thr Ala Ala Gln Arg Ala Met Met Asp
 225 230 235 240

Gln Val Arg Glu Ser Arg Asn Pro Gly Asn Ala Glu Ile Glu Asp Leu
 245 250 255

Ile Phe Leu Ala Arg Ser Ala Leu Ile Leu Arg Gly Ser Val Ala His
 260 265 270

Lys Ser Cys Leu Pro Ala Cys Val Tyr Gly Pro Ala Ile Ala Ser Gly
 275 280 285

Tyr Asn Phe Glu Lys Glu Gly Tyr Ser Leu Val Gly Ile Asp Pro Phe
 290 295 300

Lys Leu Leu Gln Asn Ser Gln Val Tyr Ser Leu Ile Arg Pro Asn Glu
 305 310 315 320

Asn Pro Ala His Lys Ser Gln Leu Val Trp Met Ala Cys Asn Ser Ala
 325 330 335

Ala Phe Glu Asp Leu Arg Val Leu Ser Phe Ile Arg Gly Thr Lys Val
 340 345 350

Ser Pro Arg Gly Lys Leu Ser Thr Arg Gly Val Gln Ile Ala Ser Asn
 355 360 365

Glu Asn Met Asp Thr Met Glu Ser Ser Thr Leu Glu Leu Arg Ser Arg
 370 375 380

Tyr Trp Ala Ile Arg Thr Arg Ser Gly Gly Asn Thr Asn Gln Gln Arg
 385 390 395 400

Ala Ser Ala Gly Gln Ile Ser Val Gln Pro Ala Phe Ser Val Gln Arg
 405 410 415

Asn Leu Pro Phe Asp Lys Pro Thr Ile Met Ala Ala Phe Thr Gly Asn
 420 425 430

Thr Glu Gly Arg Thr Ser Asp Met Arg Ala Glu Ile Ile Arg Met Met
 435 440 445

Glu Gly Ala Lys Pro Glu Glu Met Ser Phe Gln Gly Arg Gly Val Phe

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450

455

460

Glu Leu Ser Asp Glu Lys Ala Thr Asn Pro Ile Val Pro Ser Phe Asp
465 470 475 480

Met Ser Asn Glu Gly Ser Tyr Phe Phe Gly Asp Asn Ala Glu Glu Tyr
485 490 495

Asp Asn

<210> 34

<211> 469

<212> PRT

<213> Influenza A virus

<400> 34

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

Ile Ala Thr Val Cys Phe Leu Met Gln Ile Ala Ile Leu Val Thr Thr
20 25 30

Val Thr Leu His Phe Lys Gln His Glu Cys Asp Ser Pro Ala Ser Asn
35 40 45

Gln Val Met Pro Cys Glu Pro Ile Ile Ile Glu Arg Asn Ile Thr Glu
50 55 60

Ile Val Tyr Leu Asn Asn Thr Thr Ile Glu Lys Glu Ile Cys Pro Glu
65 70 75 80

Val Val Glu Tyr Arg Asn Trp Ser Lys Pro Gln Cys Gln Ile Thr Gly
85 90 95

Phe Ala Pro Phe Ser Lys Asp Asn Ser Ile Arg Leu Ser Ala Gly Gly
100 105 110

Asp Ile Trp Val Thr Arg Glu Pro Tyr Val Ser Cys Asp Pro Gly Lys
115 120 125

Cys Tyr Gln Phe Ala Leu Gly Gln Gly Thr Thr Leu Asp Asn Lys His
130 135 140

Ser Asn Asp Thr Ile His Asp Arg Ile Pro His Arg Thr Leu Leu Met
145 150 155 160

Asn Glu Leu Gly Val Pro Phe His Leu Gly Thr Arg Gln Val Cys Val
165 170 175

Ala Trp Ser Ser Ser Ser Cys His Asp Gly Lys Ala Trp Leu His Val
180 185 190

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Cys Val Thr Gly Asp Asp Lys Asn Ala Thr Ala Ser Phe Ile Tyr Asp
195 200 205
Gly Arg Leu Met Asp Ser Ile Gly Ser Trp Ser Gln Asn Ile Leu Arg
210 215 220
Thr Gln Glu Ser Glu Cys Val Cys Ile Asn Gly Thr Cys Thr Val Val
225 230 235 240
Met Thr Asp Gly Ser Ala Ser Gly Arg Ala Asp Thr Arg Ile Leu Phe
245 250 255
Ile Glu Glu Gly Lys Ile Val His Ile Ser Pro Leu Ser Gly Ser Ala
260 265 270
Gln His Val Glu Glu Cys Ser Cys Tyr Pro Arg Tyr Pro Asp Val Arg
275 280 285
Cys Ile Cys Arg Asp Asn Trp Lys Gly Ser Asn Arg Pro Val Ile Asp
290 295 300
Ile Asn Met Glu Asp Tyr Ser Ile Asp Ser Ser Tyr Val Cys Ser Gly
305 310 315 320
Leu Val Gly Asp Thr Pro Arg Asn Asp Asp Arg Ser Ser Asn Ser Asn
325 330 335
Cys Arg Asn Pro Asn Asn Glu Arg Gly Asn Pro Gly Val Lys Gly Trp
340 345 350
Ala Phe Asp Asn Gly Asp Asp Val Trp Met Gly Arg Thr Ile Ser Lys
355 360 365
Asp Leu Arg Ser Gly Tyr Glu Thr Phe Lys Val Ile Gly Gly Trp Ser
370 375 380
Thr Pro Asn Ser Lys Ser Gln Ile Asn Arg Gln Val Ile Val Asp Ser
385 390 395 400
Asn Asn Trp Ser Gly Tyr Ser Gly Ile Phe Ser Val Glu Gly Lys Arg
405 410 415
Cys Ile Asn Arg Cys Phe Tyr Val Glu Leu Ile Arg Gly Arg Gln Gln
420 425 430
Glu Thr Arg Val Trp Trp Thr Ser Asn Ser Ile Val Val Phe Cys Gly
435 440 445
Thr Ser Gly Thr Tyr Gly Thr Gly Ser Trp Pro Asp Gly Ala Asn Ile
450 455 460

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Asn Phe Met Pro Ile
465

<210> 35
<211> 121
<212> PRT
<213> Influenza A virus

<400> 35

Met Asp Ser Asn Thr Val Ser Ser Phe Gln Asp Ile Leu Leu Arg Met
1 5 10 15

Ser Lys Met Gln Leu Gly Ser Ser Ser Glu Asp Leu Asn Gly Met Ile
20 25 30

Thr Gln Phe Glu Ser Leu Lys Ile Tyr Arg Asp Ser Leu Gly Glu Ala
35 40 45

Val Met Arg Met Gly Asp Leu His Leu Leu Gln Asn Arg Asn Gly Lys
50 55 60

Trp Arg Glu Gln Leu Gly Gln Lys Phe Glu Glu Ile Arg Trp Leu Ile
65 70 75 80

Glu Glu Val Arg His Arg Leu Lys Thr Thr Glu Asn Ser Phe Glu Gln
85 90 95

Ile Thr Phe Met Gln Ala Leu Gln Leu Leu Phe Glu Val Glu Gln Glu
100 105 110

Ile Arg Thr Phe Ser Phe Gln Leu Ile
115 120

<210> 36
<211> 230
<212> PRT
<213> Influenza A virus

<400> 36

Met Asp Ser Asn Thr Val Ser Ser Phe Gln Val Asp Cys Phe Leu Trp
1 5 10 15

His Ile Arg Lys Gln Val Val Asp Gln Glu Leu Ser Asp Ala Pro Phe
20 25 30

Leu Asp Arg Leu Arg Arg Asp Gln Arg Ser Leu Arg Gly Arg Gly Asn
35 40 45

Thr Leu Gly Leu Asp Ile Lys Ala Ala Thr His Val Gly Lys Gln Ile
50 55 60

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Val Glu Lys Ile Leu Lys Glu Glu Ser Asp Glu Ala Leu Lys Met Thr
65 70 75 80

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

Glu Leu Ser Arg Asn Trp Phe Met Leu Met Pro Lys Gln Lys Val Glu
100 105 110

Gly Pro Leu Cys Ile Arg Met Asp Gln Ala Ile Met Glu Lys Asn Ile
115 120 125

Met Leu Lys Ala Asn Phe Ser Val Ile Phe Asp Arg Leu Glu Thr Ile
130 135 140

Val Leu Leu Arg Ala Phe Thr Glu Glu Gly Ala Ile Val Gly Glu Ile
145 150 155 160

Ser Pro Leu Pro Ser Phe Pro Gly His Thr Ile Glu Asp Val Lys Asn
165 170 175

Ala Ile Gly Val Leu Ile Gly Gly Leu Glu Trp Asn Asp Asn Thr Val
180 185 190

Arg Val Ser Lys Asn Leu Gln Arg Phe Ala Trp Arg Ser Ser Asn Glu
195 200 205

Asn Gly Gly Pro Pro Leu Thr Pro Lys Gln Lys Arg Lys Met Ala Arg
210 215 220

Thr Ala Arg Ser Lys Val
225 230

<210> 37
<211> 566
<212> PRT
<213> Influenza A virus

<400> 37

Met Lys Thr Ile Ile Ala Leu Ser Tyr Ile Leu Cys Leu Val Phe Ala
1 5 10 15

Gln Lys Leu Pro Gly Asn Asp Asn Ser Thr Ala Thr Leu Cys Leu Gly
20 25 30

His His Ala Val Pro Asn Gly Thr Ile Val Lys Thr Ile Thr Asn Asp
35 40 45

Gln Ile Glu Val Thr Asn Ala Thr Glu Leu Val Gln Ser Ser Ser Thr
50 55 60

Gly Gly Ile Cys Asp Ser Pro His Gln Ile Leu Asp Gly Glu Asn Cys

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65					70					75					80
Thr	Leu	Ile	Asp	Ala ₈₅	Leu	Leu	Gly	Asp	Pro ₉₀	Gln	Cys	Asp	Gly	Phe ₉₅	Gln
Asn	Lys	Lys	Trp ₁₀₀	Asp	Leu	Phe	Val	Glu ₁₀₅	Arg	Ser	Lys	Ala	Tyr ₁₁₀	Ser	Asn
Cys	Tyr	Pro ₁₁₅	Tyr	Asp	Val	Pro	Asp ₁₂₀	Tyr	Ala	Ser	Leu	Arg ₁₂₅	Ser	Leu	Val
Ala	Ser ₁₃₀	Ser	Gly	Thr	Leu	Glu ₁₃₅	Phe	Asn	Asn	Glu	Ser ₁₄₀	Phe	Asn	Trp	Thr
Gly ₁₄₅	Val	Thr	Gln	Asn	Gly ₁₅₀	Thr	Ser	Ser	Ala	Cys ₁₅₅	Lys	Arg	Arg	Ser	Asn ₁₆₀
Asn	Ser	Phe	Phe	Ser ₁₆₅	Arg	Leu	Asn	Trp	Leu ₁₇₀	Thr	His	Leu	Lys	Phe ₁₇₅	Lys
Tyr	Pro	Ala	Leu ₁₈₀	Asn	Val	Thr	Met	Pro ₁₈₅	Asn	Asn	Glu	Lys	Phe ₁₉₀	Asp	Lys
Leu	Tyr	Ile ₁₉₅	Trp	Gly	Val	His	His ₂₀₀	Pro	Gly	Thr	Asp	Asn ₂₀₅	Asp	Gln	Ile
Ser	Leu ₂₁₀	Tyr	Ala	Gln	Ala	Ser ₂₁₅	Gly	Arg	Ile	Thr	Val ₂₂₀	Ser	Thr	Lys	Arg
Ser ₂₂₅	Gln	Gln	Thr	Val	Ile ₂₃₀	Pro	Ser	Ile	Gly	Ser ₂₃₅	Arg	Pro	Arg	Ile	Arg ₂₄₀
Asp	Val	Pro	Ser	Arg ₂₄₅	Ile	Ser	Ile	Tyr	Trp ₂₅₀	Thr	Ile	Val	Lys	Pro ₂₅₅	Gly
Asp	Ile	Leu	Leu ₂₆₀	Ile	Asn	Ser	Thr	Gly ₂₆₅	Asn	Leu	Ile	Ala	Pro ₂₇₀	Arg	Gly
Tyr	Phe	Lys ₂₇₅	Ile	Arg	Ser	Gly	Lys ₂₈₀	Ser	Ser	Ile	Met	Arg ₂₈₅	Ser	Asp	Ala
Pro	Ile ₂₉₀	Gly	Lys	Cys	Asn	Ser ₂₉₅	Glu	Cys	Ile	Thr	Pro ₃₀₀	Asn	Gly	Ser	Ile
Pro ₃₀₅	Asn	Asp	Lys	Pro	Phe ₃₁₀	Gln	Asn	Val	Asn	Arg ₃₁₅	Ile	Thr	Tyr	Gly	Ala ₃₂₀
Cys	Pro	Arg	Tyr	Val ₃₂₅	Lys	Gln	Asn	Thr	Leu ₃₃₀	Lys	Leu	Ala	Thr	Gly ₃₃₅	Met
Arg	Asn	Val	Pro	Glu	Lys	Gln	Thr	Arg	Gly	Ile	Phe	Gly	Ala	Ile	Ala

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340

345

350

Gly Phe Ile Glu Asn Gly Trp Glu Gly Met Val Asp Gly Trp Tyr Gly
 355 360 365

Phe Arg His Gln Asn Ser Glu Gly Thr Gly Gln Ala Ala Asp Leu Lys
 370 375 380

Ser Thr Gln Ala Ala Ile Asn Gln Ile Asn Gly Lys Leu Asn Arg Leu
 385 390 395 400

Ile Gly Lys Thr Asn Glu Lys Phe His Gln Ile Glu Lys Glu Phe Ser
 405 410 415

Glu Val Glu Gly Arg Ile Gln Asp Leu Glu Lys Tyr Val Glu Asp Thr
 420 425 430

Lys Ile Asp Leu Trp Ser Tyr Asn Ala Glu Leu Leu Val Ala Leu Glu
 435 440 445

Asn Gln His Thr Ile Asp Leu Thr Asp Ser Glu Met Asn Lys Leu Phe
 450 455 460

Glu Arg Thr Lys Lys Gln Leu Arg Glu Asn Ala Glu Asp Met Gly Asn
 465 470 475 480

Gly Cys Phe Lys Ile Tyr His Lys Cys Asp Asn Ala Cys Ile Gly Ser
 485 490 495

Ile Arg Asn Gly Thr Tyr Asp His Asp Val Tyr Arg Asp Glu Ala Leu
 500 505 510

Asn Asn Arg Phe Gln Ile Lys Gly Val Glu Leu Lys Ser Gly Tyr Lys
 515 520 525

Asp Trp Ile Leu Trp Ile Ser Phe Ala Ile Ser Cys Phe Leu Leu Cys
 530 535 540

Val Ala Leu Leu Gly Phe Ile Met Trp Ala Cys Gln Lys Gly Asn Ile
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Arg Cys Asn Ile Cys Ile
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<210> 38

<211> 97

<212> PRT

<213> Influenza A virus

<400> 38

Met Ser Leu Leu Thr Glu Val Glu Thr Pro Ile Arg Asn Glu Trp Gly
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Cys Arg Cys Asn Asp Ser Ser Asp Pro Leu Val Val Ala Ala Ser Ile
20 25 30

Ile Gly Ile Leu His Leu Ile Leu Trp Ile Leu Asp Arg Leu Phe Phe
35 40 45

Lys Cys Val Tyr Arg Leu Phe Lys His Gly Leu Lys Arg Gly Pro Ser
50 55 60

Thr Glu Gly Val Pro Glu Ser Met Arg Glu Glu Tyr Arg Lys Glu Gln
65 70 75 80

Gln Asn Ala Val Asp Ala Asp Asp Ser His Phe Val Ser Ile Glu Leu
85 90 95

Glu

<210> 39
<211> 252
<212> PRT
<213> Influenza A virus
<400> 39

Met Ser Leu Leu Thr Glu Val Glu Thr Tyr Val Leu Ser Ile Val Pro
1 5 10 15

Ser Gly Pro Leu Lys Ala Glu Ile Ala Gln Arg Leu Glu Asp Val Phe
20 25 30

Ala Gly Lys Asn Thr Asp Leu Glu Ala Leu Met Glu Trp Leu Lys Thr
35 40 45

Arg Pro Ile Leu Ser Pro Leu Thr Lys Gly Ile Leu Gly Phe Val Phe
50 55 60

Thr Leu Thr Val Pro Ser Glu Arg Gly Leu Gln Arg Arg Arg Phe Val
65 70 75 80

Gln Asn Ala Leu Asn Gly Asn Gly Asp Pro Asn Asn Met Asp Lys Ala
85 90 95

Val Lys Leu Tyr Arg Lys Leu Lys Arg Glu Ile Thr Phe His Gly Ala
100 105 110

Lys Glu Ile Ala Leu Ser Tyr Ser Ala Gly Ala Leu Ala Ser Cys Met
115 120 125

Gly Leu Ile Tyr Asn Arg Met Gly Ala Val Thr Thr Glu Val Ala Phe
130 135 140

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Gly Leu Val Cys Ala Thr Cys Glu Gln Ile Ala Asp Ser Gln His Arg
145 150 155 160

Ser His Arg Gln Met Val Ala Thr Thr Asn Pro Leu Ile Lys His Glu
165 170 175

Asn Arg Met Val Leu Ala Ser Thr Thr Ala Lys Ala Met Glu Gln Met
180 185 190

Ala Gly Ser Ser Glu Gln Ala Ala Glu Ala Met Glu Ile Ala Ser Gln
195 200 205

Ala Arg Gln Met Val Gln Ala Met Arg Ala Val Gly Thr His Pro Ser
210 215 220

Ser Ser Thr Gly Leu Arg Asp Asp Leu Leu Glu Asn Leu Gln Thr Tyr
225 230 235 240

Gln Lys Arg Met Gly Val Gln Met Gln Arg Phe Lys
245 250

<210> 40
<211> 90
<212> PRT
<213> Influenza A virus

<400> 40

Met Glu Gln Glu Gln Asp Thr Pro Trp Thr Gln Ser Thr Glu His Thr
1 5 10 15

Asn Ile Gln Arg Arg Gly Ser Gly Arg Gln Ile Gln Lys Leu Gly His
20 25 30

Pro Asn Ser Thr Gln Leu Met Asp His Tyr Leu Arg Ile Met Ser Gln
35 40 45

Val Asp Met His Lys Gln Thr Val Ser Trp Arg Leu Trp Pro Ser Leu
50 55 60

Lys Asn Pro Thr Gln Val Ser Leu Arg Thr His Ala Leu Lys Gln Trp
65 70 75 80

Lys Ser Phe Asn Lys Gln Gly Trp Thr Asn
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<210> 41
<211> 757
<212> PRT
<213> Influenza A virus

<400> 41

Met Asp Val Asn Pro Thr Leu Leu Phe Leu Lys Val Pro Ala Gln Asn
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Ala	Ile	Ser	Thr	Thr	Phe	Pro
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				Thr		25
				Gly	Asp	Pro
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				Tyr	Ser	His
Gly	Thr	Gly	Thr	Gly	Tyr	Thr
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				Met	Asp	Thr
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				Val	Asn	Arg
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Tyr	Ser	Glu	Lys	Gly	Lys	Trp
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				Thr	Thr	Asn
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				Thr	Gly	Ala
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Gln	Leu	Asn	Pro	Ile	Asp	Gly
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				Pro	Leu	Pro
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				Glu	Asp	Asn
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				Glu	Pro	Ser
Gly	Tyr	Ala	Gln	Thr	Asp	Cys
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				Val	Leu	Glu
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				Ala	Met	Ala
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				Phe	Leu	Glu
Glu	Ser	His	Pro	Gly	Ile	Phe
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				Glu	Asn	Ser
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				Cys	Leu	Glu
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				Thr	Met	Glu
Val	Val	Gln	Gln	Thr	Arg	Val
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				Asp	Lys	Leu
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				Thr	Gln	Gly
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				Arg	Gln	Thr
Tyr	Asp	Trp	Thr	Leu	Asn	Arg
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				Asn	Gln	Pro
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				Ala	Ala	Thr
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				Ala	Leu	Ala
Asn	Thr	Ile	Glu	Val	Phe	Arg
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				Ser	Asn	Gly
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				Thr	Ala	Asn
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				Glu	Ser	Met
Gly	Arg	Leu	Ile	Asp	Phe	Leu
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				Lys	Asp	Val
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				Met	Glu	Ser
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				Met	Asp	Lys
Glu	Glu	Met	Glu	Ile	Thr	Thr
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				His	Phe	Gln
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				Arg	Lys	Arg
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				Arg	Val	Arg
Asp	Asn	Met	Thr	Lys	Lys	Met
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				Val	Thr	Gln
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				Thr	Ile	Gly
						205
				Lys	Lys	Lys
Lys	Gln	Arg	Val	Asn	Lys	Arg
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				Gly	Tyr	Leu
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				Ile	Arg	Ala
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				Leu	Lys	Arg
						230
				Arg	Arg	Ala
Asn	Thr	Met	Thr	Lys	Asp	Ala
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				Glu	Arg	Gly
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				Lys	Leu	Lys
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				Arg	Val	Tyr
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				Phe	Val	Glu
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				Gly	Phe	Val
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				Thr	Leu	Pro
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				Ser	Ile	Cys
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				Glu	Lys	Leu
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				Thr	Gln	Ser
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				Val	Arg	Lys
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				Val	Arg	Lys
						635
				Val	Arg	Lys
						640
				Val		

275

280

285

Met Met Thr Asn Ser Gln Asp Thr Glu Leu Ser Phe Thr Ile Thr Gly
 290 295 300
 Asp Asn Thr Lys Trp Asn Glu Asn Gln Asn Pro Arg Met Phe Leu Ala
 305 310 315 320
 Met Ile Thr Tyr Ile Thr Lys Asn Gln Pro Glu Trp Phe Arg Asn Ile
 325 330 335
 Leu Ser Ile Ala Pro Ile Met Phe Ser Asn Lys Met Ala Arg Leu Gly
 340 345 350
 Lys Gly Tyr Met Phe Glu Ser Lys Arg Met Lys Leu Arg Thr Gln Ile
 355 360 365
 Pro Ala Glu Met Leu Ala Ser Ile Asp Leu Lys Tyr Phe Asn Glu Ser
 370 375 380
 Thr Arg Lys Lys Ile Glu Lys Ile Arg Pro Leu Leu Ile Asp Gly Thr
 385 390 395 400
 Ala Ser Leu Ser Pro Gly Met Met Met Gly Met Phe Asn Met Leu Ser
 405 410 415
 Thr Val Leu Gly Val Ser Val Leu Asn Leu Gly Gln Lys Lys Tyr Thr
 420 425 430
 Lys Thr Thr Tyr Trp Trp Asp Gly Leu Gln Ser Ser Asp Asp Phe Ala
 435 440 445
 Leu Ile Val Asn Ala Pro Asn His Glu Gly Ile Gln Ala Gly Val Asp
 450 455 460
 Arg Phe Tyr Arg Thr Cys Lys Leu Val Gly Ile Asn Met Ser Lys Lys
 465 470 475 480
 Lys Ser Tyr Ile Asn Lys Thr Gly Thr Phe Glu Phe Thr Ser Phe Phe
 485 490 495
 Tyr Arg Tyr Gly Phe Val Ala Asn Phe Ser Met Glu Leu Pro Ser Phe
 500 505 510
 Gly Val Ser Gly Ile Asn Glu Ser Ala Asp Met Ser Ile Gly Val Thr
 515 520 525
 Val Ile Lys Asn Asn Met Ile Asn Asn Asp Leu Gly Pro Ala Thr Ala
 530 535 540
 Gln Met Ala Leu Gln Leu Phe Ile Lys Asp Tyr Arg Tyr Thr Tyr Arg

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545 550 555 560

Cys His Arg Gly Asp Thr Gln Ile Gln Thr Arg Arg Ser Phe Glu Leu
565 570 575

Lys Lys Leu Trp Asp Gln Thr Gln Ser Arg Ala Gly Leu Leu Val Ser
580 585 590

Asp Gly Gly Pro Asn Leu Tyr Asn Ile Arg Asn Leu His Ile Pro Glu
595 600 605

Val Cys Leu Lys Trp Glu Leu Met Asp Glu Asn Tyr Arg Gly Arg Leu
610 615 620

Cys Asn Pro Leu Asn Pro Phe Val Ser His Lys Glu Ile Glu Ser Val
625 630 635 640

Asn Asn Ala Val Val Met Pro Ala His Gly Pro Ala Lys Ser Met Glu
645 650 655

Tyr Asp Ala Val Ala Thr Thr His Ser Trp Asn Pro Lys Arg Asn Arg
660 665 670

Ser Ile Leu Asn Thr Ser Gln Arg Gly Ile Leu Glu Asp Glu Gln Met
675 680 685

Tyr Gln Lys Cys Cys Asn Leu Phe Glu Lys Phe Phe Pro Ser Ser Ser
690 695 700

Tyr Arg Arg Pro Ile Gly Ile Ser Ser Met Val Glu Ala Met Val Ser
705 710 715 720

Arg Ala Arg Ile Asp Ala Arg Ile Asp Phe Glu Ser Gly Arg Ile Lys
725 730 735

Lys Glu Glu Phe Ser Glu Ile Met Lys Ile Cys Ser Thr Ile Glu Glu
740 745 750

Leu Arg Arg Gln Lys
755

<210> 42
<211> 498
<212> PRT
<213> Influenza A virus

<400> 42

Met Ala Ser Gln Gly Thr Lys Arg Ser Tyr Glu Gln Met Glu Thr Asp
1 5 10 15

Gly Asp Arg Gln Asn Ala Thr Glu Ile Arg Ala Ser Val Gly Lys Met
20 25 30

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Ile Asp Gly Ile Gly Arg Phe Tyr Ile Gln Met Cys Thr Glu Leu Lys
35 40 45

Leu Ser Asp His Glu Gly Arg Leu Ile Gln Asn Ser Leu Thr Ile Glu
50 55 60

Lys Met Val Leu Ser Ala Phe Asp Glu Arg Arg Asn Lys Tyr Leu Glu
65 70 75 80

Glu His Pro Ser Ala Gly Lys Asp Pro Lys Lys Thr Gly Gly Pro Ile
85 90 95

Tyr Arg Arg Val Asp Gly Lys Trp Met Arg Glu Leu Val Leu Tyr Asp
100 105 110

Lys Glu Glu Ile Arg Arg Ile Trp Arg Gln Ala Asn Asn Gly Glu Asp
115 120 125

Ala Thr Ala Gly Leu Thr His Ile Met Ile Trp His Ser Asn Leu Asn
130 135 140

Asp Ala Thr Tyr Gln Arg Thr Arg Ala Leu Val Arg Thr Gly Met Asp
145 150 155 160

Pro Arg Met Cys Ser Leu Met Gln Gly Ser Thr Leu Pro Arg Arg Ser
165 170 175

Gly Ala Ala Gly Ala Ala Val Lys Gly Ile Gly Thr Met Val Met Glu
180 185 190

Leu Ile Arg Met Val Lys Arg Gly Ile Asn Asp Arg Asn Phe Trp Arg
195 200 205

Gly Glu Asn Gly Arg Lys Thr Arg Ser Ala Tyr Glu Arg Met Cys Asn
210 215 220

Ile Leu Lys Gly Lys Phe Gln Thr Ala Ala Gln Arg Ala Met Val Asp
225 230 235 240

Gln Val Arg Glu Ser Arg Asn Pro Gly Asn Ala Glu Ile Glu Asp Leu
245 250 255

Ile Phe Leu Ala Arg Ser Ala Leu Ile Leu Arg Gly Ser Val Ala His
260 265 270

Lys Ser Cys Leu Pro Ala Cys Ala Tyr Gly Pro Ala Val Ser Ser Gly
275 280 285

Tyr Asp Phe Glu Lys Glu Gly Tyr Ser Leu Val Gly Ile Asp Pro Phe
290 295 300

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Lys Leu Leu Gln Asn Ser Gln Ile Tyr Ser Leu Ile Arg Pro Asn Glu
305 310 315 320

Asn Pro Ala His Lys Ser Gln Leu Val Trp Met Ala Cys His Ser Ala
325 330 335

Ala Phe Glu Asp Leu Arg Leu Leu Ser Phe Ile Arg Gly Thr Lys Val
340 345 350

Ser Pro Arg Gly Lys Leu Ser Thr Arg Gly Val Gln Ile Ala Ser Asn
355 360 365

Glu Asn Met Asp Asn Met Gly Ser Ser Thr Leu Glu Leu Arg Ser Gly
370 375 380

Tyr Trp Ala Ile Arg Thr Arg Ser Gly Gly Asn Thr Asn Gln Gln Arg
385 390 395 400

Ala Ser Ala Gly Gln Thr Ser Val Gln Pro Thr Phe Ser Val Gln Arg
405 410 415

Asn Leu Pro Phe Glu Lys Ser Thr Ile Met Ala Ala Phe Thr Gly Asn
420 425 430

Thr Glu Gly Arg Thr Ser Asp Met Arg Ala Glu Ile Ile Arg Met Met
435 440 445

Glu Gly Ala Lys Pro Glu Glu Val Ser Phe Arg Gly Arg Gly Val Phe
450 455 460

Glu Leu Ser Asp Glu Lys Ala Thr Asn Pro Ile Val Pro Ser Phe Asp
465 470 475 480

Met Ser Asn Glu Gly Ser Tyr Phe Phe Gly Asp Asn Ala Glu Glu Tyr
485 490 495

Asp Asn

<210> 43
<211> 469
<212> PRT
<213> Influenza A virus

<400> 43

Met Asn Pro Asn Gln Lys Ile Ile Thr Ile Gly Ser Val Ser Leu Thr
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Ile Ser Thr Ile Cys Phe Phe Met Gln Ile Ala Ile Leu Ile Thr Thr
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Val	Thr	Leu 35	His	Phe	Lys	Gln	Tyr 40	Glu	Phe	Asn	Ser	Pro 45	Pro	Asn	Asn
Gln	Val 50	Met	Leu	Cys	Glu	Pro 55	Thr	Ile	Ile	Glu	Arg 60	Asn	Ile	Thr	Glu
Ile 65	Val	Tyr	Leu	Thr	Asn 70	Thr	Thr	Ile	Glu	Lys 75	Glu	Met	Cys	Pro	Lys 80
Leu	Ala	Glu	Tyr	Arg 85	Asn	Trp	Ser	Lys	Pro 90	Gln	Cys	Asp	Ile	Thr 95	Gly
Phe	Ala	Pro	Phe 100	Ser	Lys	Asp	Asn	Ser 105	Ile	Arg	Leu	Ser	Ala 110	Gly	Gly
Asp	Ile	Trp 115	Val	Thr	Arg	Glu	Pro 120	Tyr	Val	Ser	Cys	Asp 125	Pro	Asp	Lys
Cys	Tyr 130	Gln	Phe	Ala	Leu	Gly 135	Gln	Gly	Thr	Thr	Leu 140	Asn	Asn	Val	His
Ser 145	Asn	Asp	Thr	Val	His 150	Asp	Arg	Thr	Pro	Tyr 155	Arg	Thr	Leu	Leu	Met 160
Asn	Glu	Leu	Gly	Val 165	Pro	Phe	His	Leu	Gly 170	Thr	Lys	Gln	Val	Cys 175	Ile
Ala	Trp	Ser	Ser 180	Ser	Ser	Cys	His	Asp 185	Gly	Lys	Ala	Trp	Leu 190	His	Val
Cys	Val	Thr 195	Gly	Asp	Asp	Lys	Asn 200	Ala	Thr	Ala	Ser	Phe 205	Ile	Tyr	Asn
Gly	Arg 210	Leu	Val	Asp	Ser	Ile 215	Val	Ser	Trp	Ser	Lys 220	Lys	Ile	Leu	Arg
Thr 225	Gln	Glu	Ser	Glu	Cys 230	Val	Cys	Ile	Asn	Gly 235	Thr	Cys	Thr	Val	Val 240
Met	Thr	Asp	Gly	Ser 245	Ala	Ser	Gly	Lys	Ala 250	Asp	Thr	Lys	Ile	Leu 255	Phe
Ile	Glu	Glu	Gly 260	Lys	Ile	Ile	His	Thr 265	Ser	Thr	Leu	Ser	Gly 270	Ser	Ala
Gln	His	Val 275	Glu	Glu	Cys	Ser	Cys 280	Tyr	Pro	Arg	Tyr	Pro 285	Gly	Val	Arg
Cys	Val 290	Cys	Arg	Asp	Asn	Trp 295	Lys	Gly	Ser	Asn	Arg 300	Pro	Ile	Val	Asp

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Ile Asn Ile Lys Asp Tyr Ser Ile Val Ser Ser Tyr Val Cys Ser Gly
305 310 315 320

Leu Val Gly Asp Thr Pro Arg Lys Asn Asp Ser Ser Ser Ser Ser His
325 330 335

Cys Leu Asp Pro Asn Asn Glu Glu Gly Gly His Gly Val Lys Gly Trp
340 345 350

Ala Phe Asp Asp Gly Asn Asp Val Trp Met Gly Arg Thr Ile Ser Glu
355 360 365

Lys Leu Arg Ser Gly Tyr Glu Thr Phe Lys Val Ile Glu Gly Trp Ser
370 375 380

Lys Pro Asn Ser Lys Leu Gln Ile Asn Arg Gln Val Ile Val Asp Arg
385 390 395 400

Gly Asn Arg Ser Gly Tyr Ser Gly Ile Phe Ser Val Glu Gly Lys Ser
405 410 415

Cys Ile Asn Arg Cys Phe Tyr Val Glu Leu Ile Arg Gly Arg Lys Glu
420 425 430

Glu Thr Glu Val Leu Trp Thr Ser Asn Ser Ile Val Val Phe Cys Gly
435 440 445

Thr Ser Gly Thr Tyr Gly Thr Gly Ser Trp Pro Asp Gly Ala Asp Ile
450 455 460

Asn Leu Met Pro Ile
465

<210> 44
<211> 716
<212> PRT
<213> Influenza A virus

<400> 44

Met Glu Asp Phe Val Arg Gln Cys Phe Asn Pro Met Ile Val Glu Leu
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Ala Glu Lys Ala Met Lys Glu Tyr Gly Glu Asp Leu Lys Ile Glu Thr
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Asn Lys Phe Ala Ala Ile Cys Thr His Leu Glu Val Cys Phe Met Tyr
35 40 45

Ser Asp Phe His Phe Ile Asn Glu Gln Gly Glu Ser Ile Val Val Glu
50 55 60

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Leu Asp Asp Pro Asn Ala Leu Leu Lys His Arg Phe Glu Ile Ile Glu
 65 70 75 80
 Gly Arg Asp Arg Thr Met Ala Trp Thr Val Val Asn Ser Ile Cys Asn
 85 90 95
 Thr Thr Gly Ala Glu Lys Pro Lys Phe Leu Pro Asp Leu Tyr Asp Tyr
 100 105 110
 Lys Glu Asn Arg Phe Ile Glu Ile Gly Val Thr Arg Arg Glu Val His
 115 120 125
 Ile Tyr Tyr Leu Glu Lys Ala Asn Lys Ile Lys Ser Glu Asn Thr His
 130 135 140
 Ile His Ile Phe Ser Phe Thr Gly Glu Glu Ile Ala Thr Lys Ala Asp
 145 150 155 160
 Tyr Thr Leu Asp Glu Glu Ser Arg Ala Arg Ile Lys Thr Arg Leu Phe
 165 170 175
 Thr Ile Arg Gln Glu Met Ala Asn Arg Gly Leu Trp Asp Ser Phe Arg
 180 185 190
 Gln Ser Glu Arg Gly Glu Glu Thr Ile Glu Glu Lys Phe Glu Ile Ser
 195 200 205
 Gly Thr Met Arg Arg Leu Ala Asp Gln Ser Leu Pro Pro Lys Phe Ser
 210 215 220
 Cys Leu Glu Asn Phe Arg Ala Tyr Val Asp Gly Phe Glu Pro Asn Gly
 225 230 235 240
 Cys Ile Glu Gly Lys Leu Ser Gln Met Ser Lys Glu Val Asn Ala Lys
 245 250 255
 Ile Glu Pro Phe Leu Lys Thr Thr Pro Arg Pro Ile Lys Leu Pro Asn
 260 265 270
 Gly Pro Pro Cys Tyr Gln Arg Ser Lys Phe Leu Leu Met Asp Ala Leu
 275 280 285
 Lys Leu Ser Ile Glu Asp Pro Ser His Glu Gly Glu Gly Ile Pro Leu
 290 295 300
 Tyr Asp Ala Ile Lys Cys Ile Lys Thr Phe Phe Gly Trp Lys Glu Pro
 305 310 315 320
 Tyr Ile Val Lys Pro His Glu Lys Gly Ile Asn Ser Asn Tyr Leu Leu
 325 330 335

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Ser Trp Lys Gln Val Leu Ser Glu Leu Gln Asp Ile Glu Asn Glu Glu
340 345 350

Lys Ile Pro Arg Thr Lys Asn Met Lys Lys Thr Ser Gln Leu Lys Trp
355 360 365

Ala Leu Gly Glu Asn Met Ala Pro Glu Lys Val Asp Phe Asp Asn Cys
370 375 380

Arg Asp Ile Ser Asp Leu Lys Gln Tyr Asp Ser Asp Glu Pro Glu Leu
385 390 395 400

Arg Ser Leu Ser Ser Trp Ile Gln Asn Glu Phe Asn Lys Ala Cys Glu
405 410 415

Leu Thr Asp Ser Ile Trp Ile Glu Leu Asp Glu Ile Gly Glu Asp Val
420 425 430

Ala Pro Ile Glu Tyr Ile Ala Ser Met Arg Arg Asn Tyr Phe Thr Ala
435 440 445

Glu Val Ser His Cys Arg Ala Thr Glu Tyr Ile Met Lys Gly Val Tyr
450 455 460

Ile Asn Thr Ala Leu Leu Asn Ala Ser Cys Ala Ala Met Asp Asp Phe
465 470 475 480

Gln Leu Ile Pro Met Ile Ser Lys Cys Arg Thr Lys Glu Gly Arg Arg
485 490 495

Lys Thr Asn Leu Tyr Gly Phe Ile Ile Lys Gly Arg Ser His Leu Arg
500 505 510

Asn Asp Thr Asp Val Val Asn Phe Val Ser Met Glu Phe Ser Leu Thr
515 520 525

Asp Pro Arg Leu Glu Pro His Lys Trp Glu Lys Tyr Cys Val Leu Glu
530 535 540

Ile Gly Asp Met Leu Leu Arg Ser Ala Ile Gly Gln Ile Ser Arg Pro
545 550 555 560

Met Phe Leu Tyr Val Arg Thr Asn Gly Thr Ser Lys Val Lys Met Lys
565 570 575

Trp Gly Met Glu Met Arg Arg Cys Leu Leu Gln Ser Leu Gln Gln Ile
580 585 590

Glu Ser Met Ile Glu Ala Glu Ser Ser Ile Lys Glu Lys Asp Met Thr
595 600 605

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Lys Glu Phe Phe Glu Asn Lys Ser Glu Ala Trp Pro Ile Gly Glu Ser
610 615 620

Pro Lys Gly Val Glu Glu Gly Ser Ile Gly Lys Val Cys Arg Thr Leu
625 630 635 640

Leu Ala Lys Ser Val Phe Asn Ser Leu Tyr Ala Ser Pro Gln Leu Glu
645 650 655

Gly Phe Ser Ala Glu Ser Arg Lys Leu Leu Leu Val Val Gln Ala Leu
660 665 670

Arg Asp Asn Leu Glu Pro Gly Thr Phe Asp Leu Gly Gly Leu Tyr Glu
675 680 685

Ala Ile Glu Glu Cys Leu Ile Asn Asp Pro Trp Val Leu Leu Asn Ala
690 695 700

Ser Trp Phe Asn Ser Phe Leu Thr His Ala Leu Lys
705 710 715

<210> 45
<211> 759
<212> PRT
<213> Influenza A virus

<400> 45

Met Glu Arg Ile Lys Glu Leu Arg Asn Leu Met Ser Gln Ser Arg Thr
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Arg Glu Ile Leu Thr Lys Thr Thr Val Asp His Met Ala Ile Ile Lys
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Lys Tyr Thr Ser Gly Arg Gln Glu Lys Asn Pro Ser Leu Arg Met Lys
35 40 45

Trp Met Met Ala Met Lys Tyr Pro Ile Thr Ala Asp Lys Arg Ile Thr
50 55 60

Glu Met Val Pro Glu Arg Asn Glu Gln Gly Gln Thr Leu Trp Ser Lys
65 70 75 80

Met Ser Asp Ala Gly Ser Asp Arg Val Met Val Ser Pro Leu Ala Val
85 90 95

Thr Trp Trp Asn Arg Asn Gly Pro Val Ala Ser Thr Val His Tyr Pro
100 105 110

Lys Val Tyr Lys Thr Tyr Phe Asp Lys Val Glu Arg Leu Lys His Gly
115 120 125

Thr Phe Gly Pro Val His Phe Arg Asn Gln Val Lys Ile Arg Arg Arg

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130

135

140

Val Asp Ile Asn Pro Gly His Ala Asp Leu Ser Ala Lys Glu Ala Gln
 145 150 155 160

Asp Val Ile Met Glu Val Val Phe Pro Asn Glu Val Gly Ala Arg Ile
 165 170 175

Leu Thr Ser Glu Ser Gln Leu Thr Ile Thr Lys Glu Lys Lys Glu Glu
 180 185 190

Leu Arg Asp Cys Lys Ile Ser Pro Leu Met Val Ala Tyr Met Leu Glu
 195 200 205

Arg Glu Leu Val Arg Lys Thr Arg Phe Leu Pro Val Ala Gly Gly Thr
 210 215 220

Ser Ser Ile Tyr Ile Glu Val Leu His Leu Thr Gln Gly Thr Cys Trp
 225 230 235 240

Glu Gln Met Tyr Thr Pro Gly Gly Glu Val Arg Asn Asp Asp Val Asp
 245 250 255

Gln Ser Leu Ile Ile Ala Ala Arg Asn Ile Val Arg Arg Ala Ala Val
 260 265 270

Ser Ala Asp Pro Leu Ala Ser Leu Leu Glu Met Cys His Ser Thr Gln
 275 280 285

Ile Gly Gly Thr Arg Met Val Asp Ile Leu Arg Gln Asn Pro Thr Glu
 290 295 300

Glu Gln Ala Val Asp Ile Cys Lys Ala Ala Met Gly Leu Arg Ile Ser
 305 310 315 320

Ser Ser Phe Ser Phe Gly Gly Phe Thr Phe Lys Arg Thr Ser Gly Ser
 325 330 335

Ser Val Lys Lys Glu Glu Glu Val Leu Thr Gly Asn Leu Gln Thr Leu
 340 345 350

Lys Ile Arg Val His Glu Gly Tyr Glu Glu Phe Thr Met Val Gly Lys
 355 360 365

Arg Ala Thr Ala Ile Leu Arg Lys Ala Thr Arg Arg Leu Val Gln Leu
 370 375 380

Ile Val Ser Gly Arg Asp Glu Gln Ser Ile Ala Glu Ala Ile Ile Val
 385 390 395 400

Ala Met Val Phe Ser Gln Glu Asp Cys Met Ile Lys Ala Val Arg Gly

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405

410

415

Asp Leu Asn Phe Val Asn Arg Ala Asn Gln Arg Leu Asn Pro Met His
 420 425 430

Gln Leu Leu Arg His Phe Gln Lys Asp Ala Lys Val Leu Phe Gln Asn
 435 440 445

Trp Gly Ile Glu His Ile Asp Ser Val Met Gly Met Val Gly Val Leu
 450 455 460

Pro Asp Met Thr Pro Ser Thr Glu Met Ser Met Arg Gly Ile Arg Val
 465 470 475 480

Ser Lys Met Gly Val Asp Glu Tyr Ser Ser Thr Glu Arg Val Val Val
 485 490 495

Ser Ile Asp Arg Phe Leu Arg Val Arg Asp Gln Arg Gly Asn Val Leu
 500 505 510

Leu Ser Pro Glu Glu Val Ser Glu Thr Gln Gly Thr Glu Arg Leu Thr
 515 520 525

Ile Thr Tyr Ser Ser Ser Met Met Trp Glu Ile Asn Gly Pro Glu Ser
 530 535 540

Val Leu Val Asn Thr Tyr Gln Trp Ile Ile Arg Asn Trp Glu Ala Val
 545 550 555 560

Lys Ile Gln Trp Ser Gln Asn Pro Ala Met Leu Tyr Asn Lys Met Glu
 565 570 575

Phe Glu Pro Phe Gln Ser Leu Val Pro Lys Ala Ile Arg Ser Gln Tyr
 580 585 590

Ser Gly Phe Val Arg Thr Leu Phe Gln Gln Met Arg Asp Val Leu Gly
 595 600 605

Thr Phe Asp Thr Thr Gln Ile Ile Lys Leu Leu Pro Phe Ala Ala Ala
 610 615 620

Pro Pro Lys Gln Ser Arg Met Gln Phe Ser Ser Leu Thr Val Asn Val
 625 630 635 640

Arg Gly Ser Gly Met Arg Ile Leu Val Arg Gly Asn Ser Pro Val Phe
 645 650 655

Asn Tyr Asn Lys Thr Thr Lys Arg Leu Thr Ile Leu Gly Lys Asp Ala
 660 665 670

Gly Thr Leu Ile Glu Asp Pro Asp Glu Ser Thr Ser Gly Val Glu Ser
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675

680

685

Ala Val Leu Arg Gly Phe Leu Ile Ile Gly Lys Glu Asp Arg Arg Tyr
690 695 700

Gly Pro Ala Leu Ser Ile Asn Glu Leu Ser Asn Leu Ala Lys Gly Glu
705 710 715 720

Lys Ala Asn Val Leu Ile Gly Gln Gly Asp Val Val Leu Val Met Lys
725 730 735

Arg Lys Arg Asp Ser Ser Ile Leu Thr Asp Ser Gln Thr Ala Thr Lys
740 745 750

Arg Ile Arg Met Ala Ile Asn
755

<210> 46
<211> 580
<212> PRT
<213> CMV virus

<400> 46

Met Glu Ser Ser Ala Lys Arg Lys Met Asp Pro Asp Asn Pro Asp Glu
1 5 10 15

Gly Pro Ser Ser Lys Val Pro Arg Pro Glu Thr Pro Val Thr Lys Ala
20 25 30

Thr Thr Phe Leu Gln Thr Met Leu Arg Lys Glu Val Asn Ser Gln Leu
35 40 45

Ser Leu Gly Asp Pro Leu Phe Pro Glu Leu Ala Glu Glu Ser Leu Lys
50 55 60

Thr Phe Glu Gln Val Thr Glu Asp Cys Asn Glu Asn Pro Glu Lys Asp
65 70 75 80

Val Leu Ala Glu Leu Gly Asp Ile Leu Ala Gln Ala Val Asn His Ala
85 90 95

Gly Ile Asp Ser Ser Ser Thr Gly His Thr Leu Thr Thr His Ser Cys
100 105 110

Ser Val Ser Ser Ala Pro Leu Asn Lys Pro Thr Pro Thr Ser Val Ala
115 120 125

Val Thr Asn Thr Pro Leu Pro Gly Ala Ser Ala Thr Pro Glu Leu Ser
130 135 140

Pro Arg Lys Lys Pro Arg Lys Thr Thr Arg Pro Phe Lys Val Ile Ile
145 150 155 160

pctdk2011050460-seq1

Lys Pro Pro Val Pro Pro Ala Pro Ile Met Leu Pro Leu Ile Lys Gln
165 170 175
Glu Asp Ile Lys Pro Glu Pro Asp Phe Thr Ile Gln Tyr Arg Asn Lys
180 185 190
Ile Ile Asp Thr Ala Gly Cys Ile Val Ile Ser Asp Ser Glu Glu Glu
195 200 205
Gln Gly Glu Glu Val Glu Thr Arg Gly Ala Thr Ala Ser Ser Pro Ser
210 215 220
Thr Gly Ser Gly Thr Pro Arg Val Thr Ser Pro Thr His Pro Leu Ser
225 230 235 240
Gln Met Asn His Pro Pro Leu Pro Asp Pro Leu Ala Arg Pro Asp Glu
245 250 255
Asp Ser Ser Ser Ser Ser Ser Ser Ser Cys Ser Ser Ala Ser Asp Ser
260 265 270
Glu Ser Glu Ser Glu Glu Met Lys Cys Ser Ser Gly Gly Gly Ala Ser
275 280 285
Val Thr Ser Ser His His Gly Arg Gly Gly Phe Gly Ser Ala Ala Ser
290 295 300
Ser Ser Leu Leu Ser Cys Gly His Gln Ser Ser Gly Gly Ala Ser Thr
305 310 315 320
Gly Pro Arg Lys Lys Lys Ser Lys Arg Ile Ser Glu Leu Asp Asn Glu
325 330 335
Lys Val Arg Asn Ile Met Lys Asp Lys Asn Thr Pro Phe Cys Thr Pro
340 345 350
Asn Val Gln Thr Arg Arg Gly Arg Val Lys Ile Asp Glu Val Ser Arg
355 360 365
Met Phe Arg Asn Thr Asn Arg Ser Leu Glu Tyr Lys Asn Leu Pro Phe
370 375 380
Thr Ile Pro Ser Met His Gln Val Leu Asp Glu Ala Ile Lys Ala Cys
385 390 395 400
Lys Thr Met Gln Val Asn Asn Lys Gly Ile Gln Ile Ile Tyr Thr Arg
405 410 415
Asn His Glu Val Lys Ser Glu Val Asp Ala Val Arg Cys Arg Leu Gly
420 425 430

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Thr Met Cys Asn Leu Ala Leu Ser Thr Pro Phe Leu Met Glu His Thr
435 440 445

Met Pro Val Thr His Pro Pro Glu Val Ala Gln Arg Thr Ala Asp Ala
450 455 460

Cys Asn Glu Gly Val Lys Ala Ala Trp Ser Leu Lys Glu Leu His Thr
465 470 475 480

His Gln Leu Cys Pro Arg Ser Ser Asp Tyr Arg Asn Met Ile Ile His
485 490 495

Ala Ala Thr Pro Val Asp Leu Leu Gly Ala Leu Asn Leu Cys Leu Pro
500 505 510

Leu Met Gln Lys Phe Pro Lys Gln Val Met Val Arg Ile Phe Ser Thr
515 520 525

Asn Gln Gly Gly Phe Met Leu Pro Ile Tyr Glu Thr Ala Ala Lys Ala
530 535 540

Tyr Ala Val Gly Gln Phe Glu Gln Pro Thr Glu Thr Pro Pro Glu Asp
545 550 555 560

Leu Asp Thr Leu Ser Leu Ala Ile Glu Ala Ala Ile Gln Asp Leu Arg
565 570 575

Asn Lys Ser Gln
580

<210> 47
<211> 27
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen
<400> 47

Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Asx Gly Arg
1 5 10 15

Val Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 48
<211> 25
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

pctdk2011050460-seq1

<400> 48

Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val Ala
1 5 10 15

Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 49

<211> 27

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 49

Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Arg Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val Arg
20 25

<210> 50

<211> 26

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 50

Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 51

<211> 27

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 51

Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Arg
1 5 10 15

Val Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 52

<211> 26

<212> PRT

<213> Artificial

pctdk2011050460-seq1

<220>

<223> Artificial peptide sequence derived from antigen

<400> 52

Asx Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 53

<211> 27

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 53

Arg Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Asx Arg
1 5 10 15

Val Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 54

<211> 26

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 54

Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Lys Lys Lys Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 55

<211> 26

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 55

Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Arg Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

pctdk2011050460-seq1

<210> 56
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 56

Lys Lys Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Lys Lys Val
 1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 57
 <211> 25
 <212> PRT
 <213> Artificial

<220>
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<400> 57

Trp Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val Ala
 1 5 10 15

Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 58
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 58

Trp Trp Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val
 1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 59
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 59

Glu Glu Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Glu Glu Val
 1 5 10 15

pctdk2011050460-seq1
 Ala Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 60
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 60

Gly Gly Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Gly Gly Val
 1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 61
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 61

Glu Glu Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val
 1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 62
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 62

Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Leu Arg Arg
 1 5 10 15

Val Ala Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 63
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 63

pctdk2011050460-seq1

Trp Trp Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 64
<211> 27
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 64

Trp Trp Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Arg
1 5 10 15

Val Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 65
<211> 25
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 65

Trp Trp Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Val Ala
1 5 10 15

Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 66
<211> 25
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 66

Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val Ala
1 5 10 15

Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 67
<211> 26
<212> PRT
<213> Artificial

<220>

pctdk2011050460-seq1

<223> Artificial peptide sequence derived from antigen

<400> 67

Arg Arg Gly Tyr Leu Pro Ala Val Gly Ala Pro Ile Gly Asx Arg Val
1 5 10 15

Ile Arg Val Ile Ala His Gly Leu Arg Leu
20 25

<210> 68

<211> 26

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 68

Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Asx Arg Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 69

<211> 23

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 69

Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Gly Val Ala Arg Ala
1 5 10 15

Leu Ala His Gly Val Arg Val
20

<210> 70

<211> 25

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 70

Trp Trp Gly Tyr Leu Pro Ala Val Gly Ala Pro Ile Arg Arg Val Ile
1 5 10 15

Arg Val Ile Ala His Gly Leu Arg Leu
20 25

<210> 71

<211> 23

pctdk2011050460-seq1

<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 71

Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Gly Val Ala Arg Ala
1 5 10 15

Leu Ala His Gly Val Arg Val
20

<210> 72
<211> 27
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 72

Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Asx Gly Arg
1 5 10 15

Val Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 73
<211> 25
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 73

Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val Ala
1 5 10 15

Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 74
<211> 26
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 74

Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Arg Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

pctdk2011050460-seq1

<210> 75
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen
 <400> 75

Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val
 1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 76
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen
 <400> 76

Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Arg
 1 5 10 15

Val Ala Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 77
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen
 <400> 77

Asx Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val
 1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
 20 25

<210> 78
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen
 <400> 78

Arg Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Asx Arg
 1 5 10 15

pctdk2011050460-seq1

Val Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 79
<211> 26
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen
<400> 79

Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Lys Lys Lys Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 80
<211> 26
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen
<400> 80

Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Arg Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 81
<211> 26
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen
<400> 81

Lys Lys Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Lys Lys Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 82
<211> 25
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

pctdk2011050460-seq1

<400> 82

Trp Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val Ala
1 5 10 15

Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 83

<211> 26

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 83

Trp Trp Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Arg Arg Val
1 5 10 15

Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 84

<211> 27

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 84

Arg Arg Gly Tyr Ile Pro Leu Val Gly Ala Pro Leu Gly Leu Arg Arg
1 5 10 15

Val Ala Arg Ala Leu Ala His Gly Val Arg Val
20 25

<210> 85

<211> 24

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 85

Arg Arg Asn Tyr Val Thr Gly Asn Ile Pro Gly Asx Arg Gly Ile Thr
1 5 10 15

Phe Ser Ile Phe Leu Ile Val Ser
20

<210> 86

<211> 23

<212> PRT

<213> Artificial

pctdk2011050460-seq1

<220>

<223> Artificial peptide sequence derived from antigen

<400> 86

Trp Trp Asn Tyr Ala Thr Gly Asn Leu Pro Gly Arg Arg Cys Ser Phe
1 5 10 15

Ser Ile Phe Leu Leu Ala Leu
20

<210> 87

<211> 24

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 87

Trp Trp Asn Tyr Val Thr Gly Asn Ile Pro Gly Asx Arg Gly Ile Thr
1 5 10 15

Phe Ser Ile Phe Leu Ile Val Ser
20

<210> 88

<211> 24

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 88

Trp Trp Asn Tyr Val Thr Gly Asn Ile Pro Gly Arg Arg Gly Ile Thr
1 5 10 15

Phe Ser Ile Phe Leu Ile Val Ser
20

<210> 89

<211> 24

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 89

Arg Arg Asn Tyr Ala Thr Gly Asn Leu Pro Gly Arg Arg Gly Cys Ser
1 5 10 15

Phe Ser Ile Phe Leu Leu Ala Leu
20

pctdk2011050460-seq1

<210> 90
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 90

Arg Arg Val Thr Gly Asn Ile Pro Gly Ser Thr Tyr Ser Gly Asx Arg
 1 5 10 15

Gly Ile Thr Phe Ser Ile Tyr Leu Ile Val Ser
 20 25

<210> 91
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 91

Arg Arg Ile Arg Asn Leu Gly Arg Val Ile Glu Thr Leu Thr Gly Asx
 1 5 10 15

Arg Leu Glx Gly Tyr Ile Pro Leu Ile Gly Ala
 20 25

<210> 92
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 92

Arg Arg Ser Arg Asn Leu Gly Lys Val Ile Asp Thr Leu Thr Cys Asx
 1 5 10 15

Arg Leu Met Gly Tyr Ile Pro Leu Val Gly Ala
 20 25

<210> 93
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 93

Ser Arg Asn Leu Gly Lys Val Ile Asp Thr Leu Thr Cys Gly Phe Ala
 1 5 10 15

pctdk2011050460-seq1
 Asp Leu Met Gly Tyr Ile Pro Leu Val Gly Ala
 20 25

<210> 94
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 94

Trp Trp Ile Arg Asn Leu Gly Arg Val Ile Glu Thr Leu Thr Arg Arg
 1 5 10 15

Leu Glx Gly Tyr Ile Pro Leu Ile Gly Ala
 20 25

<210> 95
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 95

Trp Trp Ser Arg Asn Leu Gly Lys Val Ile Asp Thr Leu Thr Cys Arg
 1 5 10 15

Arg Leu Met Gly Tyr Ile Pro Leu Val Gly Ala
 20 25

<210> 96
 <211> 26
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 96

Arg Arg Gly Gly Gly Gln Ile Ile Gly Gly Asn Tyr Leu Ile Pro Arg
 1 5 10 15

Asx Pro Asx Ile Gly Val Arg Ala Thr Asx
 20 25

<210> 97
 <211> 25
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen

<400> 97

pctdk2011050460-seq1

Gly Gly Gly Gln Ile Val Gly Gly Val Tyr Leu Leu Pro Arg Arg Gly
1 5 10 15

Pro Arg Leu Gly Val Arg Ala Thr Arg
20 25

<210> 98
<211> 27
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 98

Arg Arg Gly Gly Gly Gln Ile Val Gly Gly Val Tyr Leu Leu Pro Arg
1 5 10 15

Arg Gly Pro Arg Leu Gly Val Arg Ala Thr Arg
20 25

<210> 99
<211> 26
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 99

Trp Trp Gly Gly Gly Gln Ile Val Gly Gly Val Tyr Leu Leu Pro Arg
1 5 10 15

Arg Gly Pro Arg Leu Gly Val Arg Ala Thr
20 25

<210> 100
<211> 21
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 100

Asx Arg Leu Ile Phe Leu Ala Arg Ser Ala Leu Ile Val Arg Gly Ser
1 5 10 15

Val Ala His Lys Ser
20

<210> 101
<211> 21
<212> PRT
<213> Artificial

<220>

pctdk2011050460-seq1

<223> Artificial peptide sequence derived from antigen

<400> 101

Glu Asp Leu Ile Phe Leu Ala Arg Ser Ala Leu Ile Leu Arg Gly Ser
1 5 10 15

Val Ala His Lys Ser
20

<210> 102

<211> 28

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 102

Asx Arg Leu Ile Phe Leu Ala Arg Ser Ala Leu Ile Leu Asx Gly Arg
1 5 10 15

Ser Ala Leu Ile Leu Arg Gly Ser Val Ala His Lys
20 25

<210> 103

<211> 23

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 103

Ser Ala Tyr Glu Arg Met Cys Asn Ile Leu Lys Gly Lys Phe Gln Thr
1 5 10 15

Ala Ala Gln Arg Ala Met Met
20

<210> 104

<211> 23

<212> PRT

<213> Artificial

<220>

<223> Artificial peptide sequence derived from antigen

<400> 104

Ser Ala Tyr Glu Arg Glx Val Asn Ile Leu Lys Gly Lys Phe Gln Thr
1 5 10 15

Ala Ala Gln Arg Ala Val Glx
20

<210> 105

<211> 24

pctdk2011050460-seq1

<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 105

Asx Arg Thr Ala Tyr Glu Arg Glx Cys Asn Ile Leu Asx Arg Gly Arg
1 5 10 15

Phe Gln Thr Val Val Gln Asx Ala
20

<210> 106
<211> 25
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 106

Asx Arg Ile Ala Tyr Glu Arg Met Cys Asn Ile Leu Leu Asx Arg Gly
1 5 10 15

Lys Phe Gln Thr Ala Ala Gln Arg Ala
20 25

<210> 107
<211> 21
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 107

Ile Ala Tyr Glu Arg Met Cys Asn Ile Leu Lys Gly Lys Phe Gln Thr
1 5 10 15

Ala Ala Gln Arg Ala
20

<210> 108
<211> 27
<212> PRT
<213> Artificial

<220>
<223> Artificial peptide sequence derived from antigen

<400> 108

Leu Phe Phe Lys Cys Ile Tyr Arg Leu Phe Lys His Gly Leu Lys Arg
1 5 10 15

Gly Pro Ser Thr Glu Gly Val Pro Glu Ser Met
20 25

pctdk2011050460-seq1

<210> 109
 <211> 30
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen
 <400> 109

Asx Arg Arg Leu Phe Phe Lys Thr Ile Thr Arg Leu Phe Asx His Gly
 1 5 10 15

Leu Arg Arg Leu Leu Ser Thr Glu Gly Val Pro Asn Ser Glx
 20 25 30

<210> 110
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen
 <400> 110

Asx Arg Gly Leu Glu Pro Leu Val Ile Ala Gly Ile Leu Ala Arg Arg
 1 5 10 15

Gly Ser Leu Val Gly Leu Leu His Ile Val Leu
 20 25

<210> 111
 <211> 27
 <212> PRT
 <213> Artificial

<220>
 <223> Artificial peptide sequence derived from antigen
 <400> 111

Asx Arg Gly Ser Asp Pro Leu Val Val Ala Ala Ser Ile Val Arg Arg
 1 5 10 15

Ala Ser Ile Val Gly Ile Leu His Leu Ile Leu
 20 25

<210> 112
 <211> 22
 <212> PRT
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Arg Asn Leu Val Pro Met Val Ala Thr Val Arg Arg Asn Leu Val Pro
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Met Val Ala Thr Val Asx
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Pro Met Val Ala Thr Val Asx
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Arg Asn Ile Val Pro Glx Val Val Thr Ala Arg Arg Asn Ile Val Pro
1 5 10 15

Glx Val Val Thr Ala Asx
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<211> 24
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Pro Glu Val Ile Pro Met Phe Ser Ala Leu Ser Glu Gly Ala Thr Pro
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Gln Asp Leu Asn Thr Met Leu Asn
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<400> 116

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Leu Leu Tyr Gly Ala Thr Pro Tyr Ala Ile Gly
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Gln Gly Val Gly
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Gly Asp Asp Phe Ser
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Arg Arg Gly Pro Val Val His Leu Thr Leu Arg Arg Arg Gly Gln Ala
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Gly Asp Asp Phe Ser
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Arg Arg Gly Pro Val Val His Leu Thr Leu Arg Gly Arg Arg Gly Gln
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Ala Gly Asp Asp Phe Ser
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Arg Arg Leu Glu Cys Val Tyr Cys Lys Gln Gln Leu Leu Arg Arg Glu
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Val Tyr Asp Phe Ala Phe Arg Asp Leu Cys
 20 25

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Arg Arg Gly Val Tyr Asp Phe Ala Phe Arg Asp Leu Cys Arg Arg Gly
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Phe Ala Phe Arg Asp Leu Cys Ile Val Tyr Arg
 20 25

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 <400> 123

Arg Arg Gly Val Phe Asp Tyr Ala Phe Arg Asp Ile Asn Arg Arg Gly
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Phe Ala Tyr Arg Asp Ile Asn Leu Ala Tyr Arg
20 25

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<400> 124

Arg Arg Gly Ala Thr Pro Val Asp Leu Leu Gly Ala Arg Arg Gly Ala
1 5 10 15

Leu Asn Leu Cys Leu Pro Met Arg
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<210> 125
<211> 24
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<400> 125

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Leu Gln Ile Asx Leu Pro Leu Arg
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<210> 126
<211> 98
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<400> 126

Met His Gly Asp Thr Pro Thr Leu His Glu Tyr Met Leu Asp Leu Gln
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Pro Glu Thr Thr Asp Leu Tyr Cys Tyr Glu Gln Leu Asn Asp Ser Ser
20 25 30

Glu Glu Glu Asp Glu Ile Asp Gly Pro Ala Gly Gln Ala Glu Pro Asp
35 40 45

Arg Ala His Tyr Asn Ile Val Thr Phe Cys Cys Lys Cys Asp Ser Thr
50 55 60

Leu Arg Leu Cys Val Gln Ser Thr His Val Asp Ile Arg Thr Leu Glu
65 70 75 80

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Asp Leu Leu Met Gly Thr Leu Gly Ile Val Cys Pro Ile Cys Ser Gln
85 90 95

Lys Pro

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<400> 128

Gly Tyr Leu Pro Ala Val Gly Ala Pro Ile Gly
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<400> 129

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Asn Tyr Val Thr Gly Asn Ile Pro Gly
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Asn Tyr Ala Thr Gly Asn Leu Pro Gly
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Asn Tyr Ala Thr Gly Asn Leu Pro Gly
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Val Thr Gly Asn Ile Pro Gly Ser Thr Tyr Ser
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Ser Arg Asn Leu Gly Lys Val Ile Asp Thr Leu Thr Cys
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1 5 10

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<400> 137

Gly Gly Gly Gln Ile Ile Gly Gly Asn Tyr Leu Ile Pro
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<400> 138

Gly Gly Gly Gln Ile Val Gly Gly Val Tyr Leu Leu Pro
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<400> 139

Leu Ile Phe Leu Ala Arg Ser Ala Leu Ile Val
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Leu Ile Phe Leu Ala Arg Ser Ala Leu Ile Leu
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Ser Ala Tyr Glu Arg Met Cys Asn Ile Leu
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Ser Ala Tyr Glu Arg Glx Val Asn Ile Leu
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 1 5 10

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Ile Ala Tyr Glu Arg Met Cys Asn Ile Leu

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Ile Ala Tyr Glu Arg Met Cys Asn Ile Leu
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Leu Phe Phe Lys Cys Ile Tyr Arg Leu Phe Lys His Gly Leu
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Leu Phe Phe Lys Thr Ile Thr Arg Leu Phe Asx His Gly Leu
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Gly Leu Glu Pro Leu Val Ile Ala Gly Ile Leu Ala
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Asn Leu Val Pro Met Val Ala Thr Val
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Asn Leu Val Pro Met Val Ala Thr Val
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Asn Ile Val Pro Glx Val Val Thr Ala
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Pro Glu Val Ile Pro Met Phe Ser Ala Leu Ser
1 5 10

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<220>
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<400> 156

Ala Leu Gly Pro Ala Ala Thr Leu
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<400> 157

Gly Pro Val Val His Leu Thr Leu
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Leu Glu Cys Val Tyr Cys Lys Gln Gln Leu Leu
1 5 10

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Gly Val Tyr Asp Phe Ala Phe Arg Asp Leu Cys
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Gly Val Phe Asp Tyr Ala Phe Arg Asp Ile Asn
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Gly Ala Thr Pro Val Asp Leu Leu Gly Ala
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Gly Val Thr Pro Ala Gly Leu Ile Gly Val
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Val Ala Arg Ala Leu Ala His Gly Val Arg Val
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Val Ile Arg Val Ile Ala His Gly Leu Arg Leu

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Gly Ile Thr Phe Ser Ile Phe Leu Ile Val Ser
1 5 10

<210> 166
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Cys Ser Phe Ser Ile Phe Leu Leu Ala Leu
1 5 10

<210> 167
<211> 11
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Gly Cys Ser Phe Ser Ile Phe Leu Leu Ala Leu
1 5 10

<210> 168
<211> 11
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<220>
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Gly Ile Thr Phe Ser Ile Tyr Leu Ile Val Ser
1 5 10

<210> 169
<211> 10
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Leu Glx Gly Tyr Ile Pro Leu Ile Gly Ala
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<210> 170
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Leu Met Gly Tyr Ile Pro Leu Val Gly Ala
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Pro Asx Ile Gly Val Arg Ala Thr Asx
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Gly Pro Arg Leu Gly Val Arg Ala Thr Arg
1 5 10

<210> 174
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Gly Pro Arg Leu Gly Val Arg Ala Thr
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Arg Gly Ser Val Ala His Lys Ser
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<220>

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Ser Ala Leu Ile Leu Arg Gly Ser Val Ala His Lys
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Phe Gln Thr Ala Ala Gln Arg Ala Met Met
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Gly Pro Ser Thr Glu Gly Val Pro Glu Ser Met
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Leu Leu Ser Thr Glu Gly Val Pro Asn Ser Glx
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Gly Ser Leu Val Gly Leu Leu His Ile Val Leu
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Ala Ser Ile Val Gly Ile Leu His Leu Ile Leu
1 5 10

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Asn Ile Val Pro Glx Val Val Thr Ala
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Thr Pro Gln Asp Leu Asn Thr Met Leu Asn
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<400> 188

Ala Leu Leu Tyr Gly Ala Thr Pro Tyr Ala Ile Gly
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Met Met Thr Ala Cys Gln Gly Val Gly
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Gly Gln Ala Gly Asp Asp Phe Ser
 1 5

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<400> 191

Glu Val Tyr Asp Phe Ala Phe Arg Asp Leu Cys
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<400> 192

Gly Phe Ala Phe Arg Asp Leu Cys Ile Val Tyr
 1 5 10

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Gly Phe Ala Tyr Arg Asp Ile Asn Leu Ala Tyr
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Gly Ala Leu Asn Leu Cys Leu Pro Met
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Gly Ala Leu Gln Ile Asx Leu Pro Leu
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<210> 196
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Ile Arg Val Ile Ala His Gly Leu Arg Leu Arg
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Arg Arg Ser Arg Asn Leu Gly Lys Val Ile Asp Thr Leu Thr Cys Arg
 1 5 10 15

Arg Leu Met Gly Tyr Ile Pro Leu Val Gly Ala
 20 25

<210> 198
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<220>

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<400> 198

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Ile Pro Leu Ile Gly Ala
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<210> 199

<211> 49

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<220>

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<400> 199

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1 5 10 15

Gly Tyr Ile Pro Leu Ile Gly Ala Arg Arg Ile Arg Asn Leu Gly Arg
20 25 30

Val Ile Glu Thr Leu Thr Leu Glx Gly Tyr Ile Pro Leu Ile Gly Ala
35 40 45

Arg