

CLAIMS

1. An isolated peptide consisting of an amino acid sequence of SEQ ID No. 12, or a pharmaceutical acceptable salt thereof.
2. The peptide according to claim 1, wherein said peptide has the ability to bind to an MHC class-I or -II molecule, and wherein said peptide, when bound to said MHC, is capable of being recognized by CD4 and/or CD8 T cells.
3. The peptide according to any of claims 1 to 2, wherein said peptide is modified and/or includes non-peptide bonds.
4. A fusion protein, comprising N-terminal amino acids of the HLADR antigen-associated invariant chain (Ii), and the peptide according to any of claims 1 to 3.
5. An antibody, in particular a soluble or membrane-bound antibody, preferably a monoclonal antibody or fragment thereof, that specifically recognizes SEQ ID No. 12 of the peptide according to any of claims 1 to 4, preferably the peptide according to any of claims 1 to 4 when bound to an MHC molecule.
6. A T-cell receptor, preferably soluble or membrane-bound, or a fragment thereof, that is reactive with SEQ ID No. 12 of an HLA ligand, wherein said ligand is the peptide according to any of claims 1 to 4, preferably the peptide according to any of claims 1 to 4 when bound to an MHC molecule.
7. The T-cell receptor according to claim 6, wherein said T-cell receptor is provided as a soluble molecule and optionally carries a further effector function such as an immune stimulating domain or toxin.
8. A nucleic acid, encoding for a peptide according to any one of claims 1 to 4, an antibody or fragment thereof according to claim 5, a T-cell receptor or fragment thereof according to claim 6, optionally linked to a heterologous promoter sequence, or an expression vector expressing said nucleic acid.
9. A recombinant host cell comprising the peptide according to any one of claims 1 to 4, the antibody or fragment thereof according to claim 5, the T-cell receptor or fragment thereof according to claim 6 or the nucleic acid or the expression vector according to

claim 8, wherein said host cell preferably is selected from an antigen presenting cell, such as a dendritic cell, a T cell or an NK cell.

10. An *in vitro* method for producing activated T lymphocytes, the method comprising contacting *in vitro* T cells with antigen loaded human class I or II MHC molecules expressed on the surface of a suitable antigen-presenting cell or an artificial construct mimicking an antigen-presenting cell for a period of time sufficient to activate said T cells in an antigen specific manner, wherein said antigen is a peptide according to any one of claims 1 to 4.
11. An activated T lymphocyte, produced by the method according to claim 10, that selectively recognizes SEQ ID No. 12 of a cell which presents a polypeptide comprising an amino acid sequence given in any one of claims 1 to 4.
12. A pharmaceutical composition comprising at least one active ingredient selected from the group consisting of the peptide according to any one of claims 1 to 4, the antibody or fragment thereof according to claim 5, the T-cell receptor or fragment thereof according to claim 6, the nucleic acid or the expression vector according to claim 8, the host cell according to claim 9, or the activated T lymphocyte according to claim 11, or a conjugated or labelled active ingredient, and a pharmaceutically acceptable carrier, and optionally, pharmaceutically acceptable excipients and/or stabilizers.
13. The pharmaceutical composition according to claim 12, further comprising one or more adjuvants, optionally wherein the one or more adjuvants are selected from interleukin and immunoadjuvant.
14. The pharmaceutical composition according to claim 13, wherein
 - a) the interleukin is IL-2; and/or
 - b) the immunoadjuvant is IL-15.
15. A method for producing the peptide according to any of claims 1 to 4, the antibody or fragment thereof according to claim 5, or the T-cell receptor or fragment thereof according to claim 6, the method comprising culturing the host cell according to claim 9, and isolating the peptide, the antibody or fragment thereof or the T cell receptor or fragment thereof from said host cell and/or its culture medium.

16. The peptide according to any one of claims 1 to 4, the antibody or fragment thereof according to claim 5, the T-cell receptor or fragment thereof according to claim 6, the nucleic acid or the expression vector according to claim 8, the host cell according to claim 9, or the activated T lymphocyte according to claim 11 for use in medicine.
17. Use of an effective number of activated T lymphocytes as defined in claim 11 in the manufacture of a medicament for the treatment of cancer, wherein the medicament is for killing target cells in a patient, and wherein the target cells present a polypeptide comprising an amino acid sequence given in any one of claims 1 to 4.
18. The peptide according to any one of claims 1 to 4, the antibody or fragment thereof according to claim 5, the T-cell receptor or fragment thereof according to claim 6, the nucleic acid or the expression vector according to claim 8, the host cell according to claim 9, or the activated T lymphocyte according to claim 11 for use in diagnosis and/or treatment of cancer, or for use in the manufacture of a medicament against cancer.
19. The use according to claim 17, wherein said cancer is selected from the group of ovarian cancer, hepatocellular carcinoma, colorectal carcinoma, glioblastoma, gastric cancer, esophageal cancer, non-small cell lung cancer, small cell lung cancer, pancreatic cancer, renal cell carcinoma, prostate cancer, melanoma, breast cancer, chronic lymphocytic leukemia, Non-Hodgkin lymphoma, acute myeloid leukemia, gallbladder cancer and cholangiocarcinoma, urinary bladder cancer, uterine cancer, head and neck squamous cell carcinoma, mesothelioma and other tumors that show an over-presentation of a peptide according to claim 1.
20. A kit comprising:
 - a) a container comprising a pharmaceutical composition containing the peptide according to any one of claims 1 to 4, the antibody or fragment thereof according to claim 5, the T-cell receptor or fragment thereof according to claim 6, the nucleic acid or the expression vector according to claim 8, the host cell according to claim 9, or the activated T lymphocyte according to claim 11, in solution or in lyophilized form;
 - b) optionally, a second container containing a diluent or reconstituting solution for the lyophilized formulation;
 - c) optionally, at least one more peptide selected from the group consisting of SEQ ID No. 12, and

- d) optionally, instructions for (i) use of the solution or (ii) reconstitution and/or use of the lyophilized formulation, and optionally one or more of (iii) a buffer, (iv) a diluent, (v) a filter, (vi) a needle, (vii) a syringe, or (viii) an adjuvant.
21. A method for producing a personalized anti-cancer vaccine or a compound-based and/or cellular therapy for an individual patient, said method comprising:
- a) identifying tumor-associated peptides (TUMAPs) presented by a tumor sample from said individual patient;
 - b) comparing the peptides as identified in a) with a warehouse of peptides that have been pre-screened for immunogenicity and/or over-presentation in tumors as compared to normal tissues;
 - c) selecting at least one peptide from the warehouse that matches a TUMAP identified in the patient; and
 - d) manufacturing and/or formulating the personalized vaccine or compound-based or cellular therapy based on step c);
- wherein the warehouse comprises a peptide according to SEQ ID No. 12.
22. The method according to claim 21, wherein said TUMAPs are identified by:
- a1) comparing expression data from the tumor sample to expression data from a sample of normal tissue corresponding to the tissue type of the tumor sample to identify proteins that are over-expressed or aberrantly expressed in the tumor sample; and
 - a2) correlating the expression data with sequences of MHC ligands bound to MHC class I and/or class II molecules in the tumor sample to identify MHC ligands derived from proteins over-expressed or aberrantly expressed by the tumor.
23. The method according to claim 21 or 22, wherein the sequences of MHC ligands are identified by eluting bound peptides from MHC molecules isolated from the tumor sample, and sequencing the eluted ligands.
24. The method according to any of claims 21 to 23 wherein the normal tissue corresponding to the tissue type of the tumor sample is obtained from the same patient.

25. The method according to any of claims 21 to 24, wherein the peptides included in the warehouse are identified based on the following steps:
 - aa. Performing genome-wide messenger ribonucleic acid (mRNA) expression analysis by highly parallel methods, such as microarrays or sequencing-based expression profiling, comprising identify genes that over-expressed in a malignant tissue, compared with a normal tissue or tissues;
 - ab. Selecting peptides encoded by selectively expressed or over-expressed genes as detected in step aa, and
 - ac. Determining an induction of *in vivo* T-cell responses by the peptides as selected comprising *in vitro* immunogenicity assays using human T cells from healthy donors or said patient; or
 - ba. Identifying HLA ligands from said tumor sample using mass spectrometry;
 - bb. Performing genome-wide messenger ribonucleic acid (mRNA) expression analysis by highly parallel methods, such as microarrays or sequencing-based expression profiling, comprising identify genes that over-expressed in a malignant tissue, compared with a normal tissue or tissues;
 - bc. Comparing the identified HLA ligands to said gene expression data;
 - bd. Selecting peptides encoded by selectively expressed or over-expressed genes as detected in step bc;
 - be. Re-detecting of selected TUMAPs from step bd on tumor tissue and lack of or infrequent detection on healthy tissues and confirming the relevance of over-expression at the mRNA level; and
 - bf. Determining an induction of *in vivo* T-cell responses by the peptides as selected comprising *in vitro* immunogenicity assays using human T cells from healthy donors or said patient.
26. The method according to any of claims 21 to 25, wherein the immunogenicity of the peptides included in the warehouse is determined by a method comprising *in vitro* immunogenicity assays, patient immunomonitoring for individual HLA binding, MHC multimer staining, ELISPOT assays and/or intracellular cytokine staining.
27. The method according to any of claims 21 to 26, further comprising identifying at least one mutation that is unique to the tumor sample relative to normal corresponding

tissue from the individual patient, and selecting a peptide that correlates with the mutation for inclusion in the vaccine or for the generation of cellular therapies.

28. The method according to claim 27, wherein said at least one mutation is identified by whole genome sequencing.