186 BD1 CIP FWC III ROÖ IN THE UNITED STATES PATENT AND TRADEMARK OFFICE Applicants : Sherie L. Morrison, et al. rial No. 07/893,610 RADE Filed June 3, 1992 For RECEPTORS BY DNA SPLICING AND EXPRESSION Art Unit 1806 T. Nisbet Examiner August 23, 1993 Hon. Commissioner of Patents and Trademarks Washington, D.C. 20231 RESPONSE UNDER 37 C.F.R. §1.115 TO EXAMINER'S ACTION Sir: Applicants respectfully request entry of the following amendments to the pending claims: **...** 3/Â (Thrice amended) A method for producing a functional antibody having a heavy chain and a light chain [two subunits] λ which comprises the steps of: transfecting a non-antibody producing 1 m DNOIL mammalian [lymphoud] cell with a first DNA sequence coding for a first <u>chain</u> (subunit) of the antibody; (b) transfecting the cell with a second DNA sequence, said second DNA sequence coding for a second chain [subunit] of the antibody, said second <u>chain</u> [subunit] being a <u>chain</u> [subunit] other than the first <u>chain</u> [subunit] <u>and</u> <u>said first and second chains being either the heavy chain or</u> the light chain; and (C) maintaining the cell in a nutrient medium, so that the cell expresses the first and second DNA sequences and the resultant chains [subunits] are intracellularly assembled together to form the antibody which is then

a hoth capable of specifically binding to secreted antigen 4,5 (Twice Amended) A method as recited in claim <u>39</u> [43] wherein the cell does not endogenously produce any immunoglobulin chains. (Amended) A method as recited in claim 39 [43] wherein the cell endogenously produces [only] an immunoglobulin light chain or an immunoglobulin heavy chain but not both 154. (Twice amended) A method for producing a functional antibody having <u>a heavy chain and a light chain</u> [two subunits], which comprises the steps of: transfecting a non-antibody producing (a) mammalian [lymphoid] cell with a plasmid comprising a first DNA sequence coding for a first chain [subunit] \Im ϕ f the antibody and a second DNA sequence coding for a second <u>chain</u> [subunit] of the antibody, said second chain [subunit] beind a chain [subunit] other than the first chain [subunit] and said first and second chains being either the heavy chain or the light chain; and maintaining the cell in a nutrient medium so (b) that the cell expresses said first DNA sequence and said second DNA sequence and the resultant chains [subunits] are intracellularly assembled together to form the antibody which is then secreted in a form capable of specifically binding to antigen.

:C Ľ Please add the following claims: A) method as recited in claim 39 wherein the cell is a lymphoid cell. 57. A method as recited in claim 54 wherein the cell is transfected via protoplast fusion. $\sqrt{2}$ 58. A method as recited in claim 54 wherein the cell is transfected via calcium phosphate precipitation. as recited in claim 54 wherein the A method cell is a lymphoid co a method as recited in claim 54 wherein the 60. cell is a myeloma cell. 61. A method as regited in claim 60 wherein the 17 1. ... cell is a murine myeloma cell. A method as revited in claim 54 wherein the 62. 1B cell does not endogenously produce any immunoglobulin chains. 63. A method as recited in claim 62 wherein the 1 4. cell is a murine P, cell 64. A method as recited in claim 54 wherein the cell endogenously produces an immunoglobulin light chain or an immunoglobulin heavy chain but not both.

7 21 65. A method as recited in claim 64 wherein the cell is a murine J558L cell.

A method as recited in claim 54 wherein the antibody is a chimeric antibody having a variable region substantially the same as that found in a first mammalian source and having a constant region substantially the same as that found in a second mammalian source, said second mammalian source being from a mammalian species other than that of the first mammalian source.

 γ^{67} . A method for producing a functional antibody having a heavy chain and a light chain which comprises the steps of:

(a) maintaining in a nutrient medium a nonantibody producing mammalian dell, said cell having been transfected with a first DNA sequence coding for a first chain of the antibody and a second DNA sequence coding for a second chain of the antibody said second chain being a chain other than the first chain and said first and second chains being either the heavy chain or the light chain;

(b) expressing from said cell the heavy chain and the light chain functionally assembled together to form said antibody which is then secreted in a form capable of binding antigen; and

(c) recovering said antibody.

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24 68. A method as recited in claim 67 wherein the cell is transfected via protoplast fusion.

cell is transfected via calcium phosphate precipitation.

method as recited in claim 67 wherein the 70. A cell is a lymphoid sell. A method as recited in claim 67 wherein the 71. <u>ب</u> ۲۰ cell is a myeloma cel $m\lambda$. A method as recited in claim 71 wherein the 12 5 cell is a murine myeloma dell. A method as recited in claim 67 wherein the cell does not endogenously produce any immunogiobulin chains. 74. A method as recited in claim 73 wherein the cell is a murine P. cell. - > 75. A method as recited in claim 67 wherein the cell endogenously produces an immenoglobulin light chain or an immunoglobulin heavy chain but not both. 76. A method as recited in claim 75 wherein the 1 cell is a murine J558L cell. 70. (A method as recited in claim 67 wherein the antibody is a chimeric antibody having a variable region substantially the same as that found in a first mammalian source and having a constant region substantially the same as that found in a second mammaliah source, said second , mammalian source being from a mammalian species other than that of the first mammalian source. Please cancel claims 49-51.

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