

# (12) International Application Status Report

**Received at International Bureau:** 12 April 2020 (12.04.2020)

**Information valid as of:** 07 September 2020 (07.09.2020)

**Report generated on:** 27 January 2021 (27.01.2021)

**(10) Publication number:**

WO2020/198683

**(43) Publication date:**

01 October 2020 (01.10.2020)

**(26) Publication language:**

English (EN)

**(21) Application Number:**

PCT/US2020/025469

**(22) Filing Date:**

27 March 2020 (27.03.2020)

**(25) Filing language:**

English (EN)

**(31) Priority number(s):**

62/825,726 (US)

**(31) Priority date(s):**

28 March 2019 (28.03.2019)

**(31) Priority status:**

Priority document received (in compliance with PCT Rule 17.1)

**(51) International Patent Classification:**

A61K 31/5377 (2006.01)

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**(54) Title (EN):** HETEROMULTIMERIC PROTEINS AND METHODS OF USE THEREOF

**(54) Title (FR):** PROTÉINES HÉTÉROMULTIMÈRES ET LEURS MÉTHODES D'UTILISATION

**(57) Abstract:**

**(EN):** The present application provides heteromultimeric proteins, such as bispecific antibodies, comprising a first antibody heavy chain constant domain 3 (CH3)-containing polypeptide having a substitution relative to a wildtype CH3 domain at amino acid position 354 with a bulky hydrophobic amino acid, and/or a second CH3 -containing polypeptide comprising a substitution relative to a wildtype CH3 domain at amino acid position 347 with a negatively charged amino acid residue. Also provided are polypeptides, nucleic acids and vectors encoding such polypeptides, pharmaceutical compositions, methods of preparation and methods of treatment using the heteromultimeric proteins.

**(FR):** La présente invention concerne des protéines hétéromultimères, telles que des anticorps bispécifiques, comprenant un premier polypeptide contenant un domaine 3 constant de chaîne lourde (CH3) d'anticorps, comportant une substitution par rapport à un domaine CH3 de type sauvage, à la position d'acide aminé 354, avec un acide aminé hydrophobe volumineux, et/ou un second polypeptide contenant CH3 comprenant une substitution par rapport à un domaine CH3 de type sauvage, à la position d'acide aminé 347, avec un résidu d'acide aminé chargé négativement. L'invention concerne également des polypeptides, des acides nucléiques et des vecteurs codant pour de tels polypeptides, des compositions pharmaceutiques, des procédés de préparation et des méthodes de traitement utilisant ces protéines hétéromultimères.

**International search report:**

Received at International Bureau: 05 September 2020 (05.09.2020) [US]

**International Report on Patentability (IPRP) Chapter II of the PCT:**

Not available

**(81) Designated States:**

AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CZ, DE, DJ, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IR, IS, JO, JP, KE, KG, KH, KN, KP, KR, KW, KZ, LA, LC, LK, LR, LS, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC, SD, SE, SG, SK, SL, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, WS, ZA, ZM, ZW

European Patent Office (EPO) : AL, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MK, MT, NL, NO, PL, PT, RO, RS, SE, SI, SK, SM, TR

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Eurasian Patent Organization (EAPO) : AM, AZ, BY, KG, KZ, RU, TJ, TM