

(19) World Intellectual Property Organization
International Bureau(43) International Publication Date
8 February 2007 (08.02.2007)

PCT

(10) International Publication Number
WO 2007/015757 A3

(51) International Patent Classification:

C07C 233/58 (2006.01) **A61P 35/00** (2006.01)
A61K 31/16 (2006.01) **C07C 49/557** (2006.01)(74) Agent: **HART, Daniel**; KNOBBE MARTENS OLSON & BEAR LLP, 2040 Main Street, 14th Floor, Irvine, California 92614 (US).

(21) International Application Number:

PCT/US2006/027385

(22) International Filing Date: 14 July 2006 (14.07.2006)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:

60/701,932 21 July 2005 (21.07.2005) US
60/734,590 7 November 2005 (07.11.2005) US
60/734,679 7 November 2005 (07.11.2005) US
60/749,542 12 December 2005 (12.12.2005) US
60/785,223 22 March 2006 (22.03.2006) US(71) Applicant (for all designated States except US): **NEREUS PHARMACEUTICALS, INC.** [US/US]; 10480 Wateridge Circle, San Diego, California 92121 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): **PALLADINO, Michael A.** [US/US]; 3357 Fortuna Ranch Road, Olivenhain, California 92024 (US). **THEODORAKIS, Emmanuel A.** [US/US]; 3905 Caminito Terviso, San Diego, California 92122 (US). **MACHERLA, Venkat Rami Reddy** [IN/US]; 10868 Caminito Arcada, San Diego, California 92131 (US). **CHAO, Ta-Hsiang** [US/US]; 7016 Enders Avenue, San Diego, California 92122 (US). **SUH, Young Ger** [KR/KR]; 1223-602, Mokryun Apt., Sanbon-dong, Gunpo, Kyungki-do 435-040 (KR).

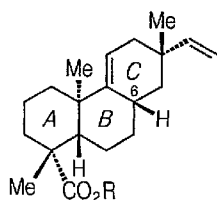
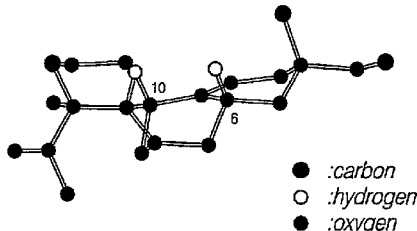
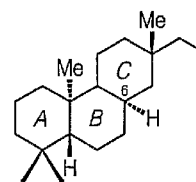
(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

— with international search report
— before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments(88) Date of publication of the international search report:
21 June 2007

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: INTERLEUKIN-1 AND TUMOR NECROSIS FACTOR- α MODULATORS; SYNTHESIS OF SUCH MODULATORS AND METHODS OF USING SUCH MODULATORS1: R=H; acanthoic acid
2: R=Me; acanthoic acid methyl esterStereochemical view of acanthoic acid (1)
(Model minimized using Chem3D)

3

Skeletal type of pimaranes
(relative stereochemistry)(57) Abstract: Compounds are disclosed that have the chemical structure of Formula (II), (IIA) and (IIB) and their prodrug esters and acid-addition salts, and that are useful as Interleukin-1 and Tumor Necrosis Factor- α modulators, and thus are useful in the treatment of various diseases, wherein the R groups are defined in the claims.