



(51) International Patent Classification:

A61K 31/40 (2006.01) C07D 409/04 (2006.01)
C07D 401/06 (2006.01) C07D 207/16 (2006.01)

(21) International Application Number:

PCT/EP2013/060834

(22) International Filing Date:

27 May 2013 (27.05.2013)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

61/653,133 30 May 2012 (30.05.2012) US

(71) Applicant (for all designated States except US): **F. HOFFMANN-LA ROCHE AG** [CH/CH]; Grenzacherstrasse 124, CH-4070 Basel (CH).

(71) Applicant (for US only): **HOFFMANN-LA ROCHE INC.** [US/US]; 340 Kingsland Street, Nutley, New Jersey 07110 (US).

(72) Inventors: **BARTKOVITZ, David Joseph**; 21 Povershon Road, Nutley, New Jersey 07110 (US). **CHU, Xin-Jie**; Room 601, No. 21, Lane 728 Changdao Road, Pudong District, Shanghai 200129 (CN). **EHRLICH, George**; 5450 Mosholu Ave., Bronx, New York 10471 (US). **LIU, Jin-Jun**; 50 Highmount Avenue, Warren, New Jersey 07059 (US). **MICHEL, Hanspeter**; 31 Perry Road, Bloomfield, New Jersey 07003 (US). **VU, Binh Thanh**; 1 Willow Lane, North Caldwell, New Jersey 07006 (US).

ZHAO, Chunlin; 590 Ruiqing Road, Zhangjiang East Campus, Pudong, Shanghai 201201 (CN).

(74) Agent: **BEYERMANN, Jochen**; Grenzacherstrasse 124, CH-4070 Basel (CH).

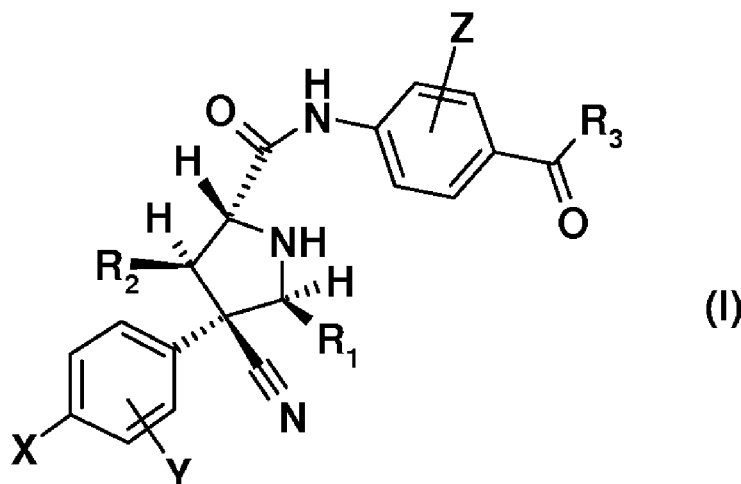
(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, 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, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, 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, LR, LS, MW, MZ, NA, RW, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (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), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, KM, ML, MR, NE, SN, TD, TG).

Published:

— with international search report (Art. 21(3))

(54) Title: SUBSTITUTED PYRROLIDINE-2-CARBOXAMIDES



(57) Abstract: There are provided compounds of the formula (I) wherein X, Y, Z, R₁, R₂ and R₃ are as described herein, enantiomers and pharmaceutically acceptable salts and esters thereof, methods of making those compounds, as well as the use of such compounds as medicaments, especially as anticancer agents.