(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization
International Bureau

(43) International Publication Date
7 August 2003 (07.08.2003)

PCT

(10) International Publication Number
WO 03/064413 A1

(51) International Patent Classification[7]: C07D 403/12, 471/04, 401/14, 491/04, A61K 31/517, A61P 35/00

(21) International Application Number: PCT/GB03/00343

(22) International Filing Date: 28 January 2003 (28.01.2003)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:
02290242.3 1 February 2002 (01.02.2002) EP

(71) Applicant *(for AE, AG, AL, AM, AT, AU, AZ, BA, BB, BE, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CY, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, FR, GB, GD, GE, GH, GM, GR, HR, HU, ID, IE, IL, IN, IS, IT, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MC, MD, MK, MN, MW, MX, MZ, NL, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SZ, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW only)*: **ASTRAZENECA R & D SODERTALJE** [SE/SE]; Sodertalje, S-151 85 (SE).

(71) Applicant *(for MG only)*: **ASTRAZENECA UK LIMITED** [GB/GB]; 15 Stanhope Gate, London, Greater London W1K 1LN (GB).

(72) Inventor; and
(75) Inventor/Applicant *(for US only)*: **HENNEQUIN, Laurent, Francois, Andre** [FR/GB]; AstraZeneca R & D Alderley, Alderley Park, Macclesfield, Cheshire SK10 4TG (GB).

(74) Agent: **ASTRAZENECA**; Global Intellectual Property, Mereside, Alderley Park, Macclesfield, Cheshire SK10 4TG (GB).

(81) Designated States *(national)*: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.

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

Published:
— *with international search report*

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.

WO 03/064413 A1

(54) Title: QUINAZOLINE COMPOUNDS

(57) Abstract: The invention relates to compounds of the formula (I): wherein ring C is as defined herein, for example indolyl, indazolyl or azaindolyl; Z is -O-, -NH- or -S-; n is 0-5; m is 0-3; R^1 and R^2 are defined herein including groups: (i) Q^1X^1 wherein Q^1 and X^1 are as defined herein; (ii) $Q^{15}W^3$ wherein Q^{15} and W^3 are as defined herein, (iii) $Q^{21}W^4C_{1-5}alkylX^1$- wherein Q^{21}, W^4 and X^1 are as defined herein, (iv) $Q^{28}C_{1-5}alkylX^1$-, $Q^{28}C_{2-5}alkenylX^1$- or $Q^{28}C_{2-5}alkynylX^1$- wherein Q^{28} and X^1 are as defined herein and (v) $Q^{29}C_{1-5}alkylX^1$-, $Q^{29}C_{2-5}alkenylX^1$- or $Q^{29}C_{2-5}alkynylX^1$- wherein Q^{29} and X^1 are as defined herein; R^2 can also be 6,7-methylenedioxy or 6,7-ethylenedioxy;and salts thereof; their use in the manufacture of a medicament for use in the production of an antiangiogenic and/or vascular permeability reducing effect in warm?blooded animals; processes for the preparation of such compounds; intermediates used in such processes; processes for making such intermediates; pharmaceutical compositions containing a compound of formula I or a pharmaceutically acceptable salt thereof and methods of treating disease states involving angiogenesis by administering a compound of formula I or a pharmaceutically acceptable salt thereof. The compounds of formula I inhibit the effects of VEGF, a property of value in the treatment of a number of disease states including cancer and rheumatoid arthritis.