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

(19) World Intellectual Property Organization
International Bureau

(43) International Publication Date
18 May 2006 (18.05.2006)

PCT

(10) International Publication Number
WO 2006/052128 A1

(51) International Patent Classification:
A61K 38/17 (2006.01) *A61P 35/00* (2006.01)
A61K 38/55 (2006.01)

(21) International Application Number:
PCT/NL2005/000788

(22) International Filing Date:
10 November 2005 (10.11.2005)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:
04078090.0 10 November 2004 (10.11.2004) EP
05075806.9 6 April 2005 (06.04.2005) EP

(71) Applicant *(for all designated States except US)*: **Hubrecht Laboratorium** [NL/NL]; Uppsalalaan 8, NL-3584 CT Utrecht (NL).

(72) Inventors; and

(75) Inventors/Applicants *(for US only)*: **CLEVERS, Johannes Carolus** [NL/NL]; Ruysdaellaan 7, NL-3712 AP Huis ter Heide (NL). **VAN GIJN, Maria, Elisabeth** [NL/NL]; Vechtensteinlaan 37a, NL-3601 CM Maarssen (NL). **VAN ES, Johannes, Hendrikus** [NL/NL]; Vechtensteinlaa 37a, NL-3601 CM Maarssen (NL).

(74) Agent: **WINCKELS, J.H.F.**; Johan de Wittlaan 7, NL-DEN HAAG 2517 JR (NL).

(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, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, 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, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, 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*

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:** TREATMENT OF AN INTESTINAL ADENOMA AND/OR ADENOCARCINOMA BY INHIBITION OF NOTCH PATHWAY ACTIVATION

(57) **Abstract:** The invention relates to the field of biochemistry and medicine. More specifically the invention relates to methods and pharmaceuticals for the treatment of intestinal neoplasia. The invention discloses the surprising finding that inhibitors of Notch pathway activation (for example g-secretase inhibitors) are extremely useful in the treatment of intestinal adenoma and/or adenocarcinoma

WO 2006/052128 A1