(19) World Intellectual Property Organization

International Bureau





(43) International Publication Date 28 February 2008 (28.02.2008)

(10) International Publication Number WO 2008/023333 A3

- (51) International Patent Classification: C07D 215/42 (2006.01)
- (21) International Application Number:

PCT/IB2007/053334

- (22) International Filing Date: 21 August 2007 (21.08.2007)
- (25) Filing Language: English
- (26) Publication Language: English
- (30) Priority Data:

1886/DEL/2006 22 August 2006 (22.08.2006) IN

- (71) Applicant (for all designated States except US): RAN-BAXY LABORATORIES LIMITED [IN/IN]; Plot No. 90, Sector - 32, Gurgaon, Haryana 122001 (IN).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): JAYACHANDRA, Suresh, Babu [IN/IN]; C-237, Phase - I, Sushant Lok, Gurgaon, Haryana 122001 (IN). SHAH, Jigar, Bhaskarbhai [IN/IN]; B-753, Tumdi Street, Sonari Mahal, Bharuch, Gujarat 392001 (IN). CHIGURU, Sailu [IN/IN]; s/o Laxman Chiguru, Kondapoor, Musthabad, Karimnager, Andhra Pradesh 505303 (IN). KHANDURI, Chandra, Has [IN/IN]; D-1952, Palam Vihar, Gurgaon, Haryana 122017 (IN).
- (74) Common Representative: RANBAXY LABORATO-RIES LIMITED; C/o DESHMUKH, Jay, R., 600 College Road East, Suite 2100, Princeton, NJ 08540 (US).

- (81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, 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, KM, 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, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, 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, MT, 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: 15 May 2008

(54) Title: PROCESS FOR THE PREPARATION OF 4-(ISOBUTYLAMINO)-3-AMINO-QUINOLINE

(57) Abstract: The invention relates to a process for the preparation of 4-isobutylamino-3- aminoquinoline. This compound is a useful pharmaceutical intermediate for the preparation of 1-(2-methylpropyl)-1H-imidazo[4,5-c]quinoline-4-amine - a compound known generically as imiquimod, an immune response modifier.

INTERNATIONAL SEARCH REPORT

International application No PCT/IB2007/053334

A. CLASSIFICATION OF SUBJECT MATTER INV. C07D215/42

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

 $\begin{tabular}{ll} Minimum documentation searched (classification system followed by classification symbols) \\ CO7D \end{tabular}$

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, BEILSTEIN Data, WPI Data

| Category* | Citation of document, with indication, where appropriate, of the | Relevant to claim No. | |
|--|--|--|--|
| Y | US 4 689 338 A (GERSTER JOHN F 25 August 1987 (1987-08-25) cited in the application column 4, line 22 - column 6, example 16 column 10; table II | 1-10 | |
| Y | US 4 988 815 A (ANDRE JEAN-DEN AL) 29 January 1991 (1991-01-2 column 6, line 54 - column 7, example 13 | 1-10 | |
| | | | |
| | | | |
| X Furt | her documents are listed in the continuation of Box C. | X See patent family annex. | |
| * Special of "A" docume consider filling of "L" docume which citatio "O" docume other "P" docume | categories of cited documents : ent defining the general state of the art which is not dered to be of particular relevance document but published on or after the international | "T" later document published after the iron priority date and not in conflict worked to understand the principle or invention "X" document of particular relevance; the cannot be considered novel or can involve an inventive step when the "Y" document of particular relevance; the cannot be considered to involve an document is combined with one or ments, such combination being obtain the art. "&" document member of the same pater | ith the application but theory underlying the seclaimed invention not be considered to document is taken alone seclaimed invention inventive step when the more other such docuvious to a person skilled |
| * Special of Commercial Commercia | categories of cited documents: ent defining the general state of the art which is not dered to be of particular relevance document but published on or after the international date ent which may throw doubts on priority claim(s) or is cited to establish the publication date of another n or other special reason (as specified) ent referring to an oral disclosure, use, exhibition or means ent published prior to the international filing date but | "T" later document published after the interpretation or priority date and not in conflict with cited to understand the principle or invention "X" document of particular relevance; the cannot be considered novel or can involve an inventive step when the "Y" document of particular relevance; the cannot be considered to involve an document is combined with one or ments, such combination being obtain the art. | ith the application but theory underlying the sectaimed invention not be considered to document is taken alone sectaimed invention inventive step when the more other such docuvious to a person skilled |
| * Special of the consideration of the country of th | categories of cited documents: ent defining the general state of the art which is not dered to be of particular relevance document but published on or after the international date ent which may throw doubts on priority claim(s) or is cited to establish the publication date of another n or other special reason (as specified) ent referring to an oral disclosure, use, exhibition or means ent published prior to the international filing date but than the priority date claimed | "T" later document published after the incomposition or priority date and not in conflict work cited to understand the principle or invention "X" document of particular relevance; the cannot be considered novel or can involve an inventive step when the "Y" document of particular relevance; the cannot be considered to involve an document is combined with one or ments, such combination being obtain the art. "&" document member of the same pater. | ith the application but theory underlying the sectaimed invention not be considered to document is taken alone sectaimed invention inventive step when the more other such docuvious to a person skilled |

INTERNATIONAL SEARCH REPORT

International application No
PCT/IB2007/053334

| C(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT | | | | | | |
|--|--|-----------------------|--|--|--|--|
| ategory* | Citation of document, with indication, where appropriate, of the relevant passages | Relevant to claim No. | | | | |
| (| JOHN F. GERSTER ET AL: "Synthesis and Structure - Activity - Relationships of 1H-Imidazo[4,5-c]quinolines that Induce Interferon Production" JOURNAL OF MEDICINAL CHEMISTRY, vol. 48, 2005, pages 3481-3491, XP002450996 cited in the application page 3488; example 12 scheme 2 | 1-10 | | | | |
| , | RICHARD C. LAROCK: "Comprehensive Organic Transformations" 1999, WILEY-VCH , XP002469288 page 821 - page 828 | 1-10 | | | | |
| Y | S. VANGAPANDU ET AL: "Ring-substituted quinolines as potential anti-tuberculosis agents" BIOORGANIC & MEDICINAL CHEMISTRY, vol. 12, 2004, pages 2501-2508, XP002469287 page 2502 scheme 1 | 1-10 | | | | |
| Y | US 2003/055087 A1 (SHINKAI HISASHI [JP] ET AL) 20 March 2003 (2003-03-20) paragraph [0206] paragraph [0216] - paragraph [0218] | 1-10 | | | | |

INTERNATIONAL SEARCH REPORT

International application No
PCT/IB2007/053334

| Patent document cited in search report | | Publication date | | Patent family member(s) | Publication date |
|---|----|---------------------|------|-------------------------|------------------|
| US 4689338 | Α | 25-08-1987 | CZ | 9103904 A3 | 17-03-1993 |
| | | | MΧ | 9203474 A1 | 01-07-1992 |
| | | | SK | 390491 A3 | 12-09-2000 |
| US 4988815 | Α | 29-01-1991 | AT | 126790 T | 15-09-1995 |
| | | | AU | 641693 B2 | 30-09-1993 |
| | | | AU | 6392890 A | 04-07-1991 |
| | | | BR | 9005452 A | 17-09-1991 |
| | | | CA | 2027245 A1 | 27-04-1991 |
| | | | DE | 69021826 D1 | 28-09-1995 |
| | | | DE | 69021826 T2 | 18-04-1996 |
| | | | EP | 0425306 A2 | 02-05-1991 |
| | | | ES | 2075168 T3 | 01-10-1995 |
| | | | HK | 1006569 A1 | 05-03-1999 |
| | | | HU | 55777 A2 | 28-06-1991 |
| | | | ΙE | 903597 A1 | 08-05-1991 |
| | | | JP | 2945121 B2 | 06-09-1999 |
| | | | JP | 3206078 A | 09-09-1991 |
| | | | NO | 904625 A | 29-04-1991 |
| | | | RU | 2083563 C1 | 10-07-1997 |
| | | | ZA | 9008193 A | 28-08-1991 |
| US 2003055087 | A1 | 20-03-2003 | NONE | | |