26/09/02
200205896 4

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

(19) World Intellectual Property Organization
International Bureau

(43) International Publication Date
11 October 2001 (11.10.2001)

PCT

(10) International Publication Number
WO 01/74329 A3

(51) International Patent Classification[7]: A61K 31/52, 9/16, 9/50

(21) International Application Number: PCT/US01/10078

(22) International Filing Date: 29 March 2001 (29.03.2001)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:
60/193,588 30 March 2000 (30.03.2000) US

(71) Applicant: **BRISTOL-MYERS SQUIBB COMPANY** [US/US]; P.O. Box 4000, Princeton, NJ 08543-4000 (US).

(72) Inventors: **ABRAMOWITZ, Rober**; 4 Abbington Lane, West Windsor, NJ 08550 (US). **O'DONOGHUE, Denise, M.**; 163 Mechanic Street, Hightstown, NJ 08520 (US). **JAIN, Nemichand, B.**; 19 Abbington Lane, West Windsor, NJ 08550 (US).

(74) Agent: **MCNEIL, Scott, Alexander**; Bristol-Myers Squibb Company, 5 Research Parkway, P.O. Box 5100, Wallingford, CT 06492-7660 (US).

(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, 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, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW.

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

Published:
— *with international search report*

(88) Date of publication of the international search report:
10 May 2002

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.

162162

ACTION

WO 01/74329 A3

(54) Title: SUSTAINED RELEASE BEADLETS CONTAINING STAVUDINE

(57) Abstract: Extended dosage forms of stavudine are provided comprising beadlets formed by extrusion-spheronization and coated with a seal coating. The beadlets are also coated with a modified release coating such that a hard gelatin capsule containing such beadlets will provide blood levels of stavudine over approximately 24 hours. The beadlets are prepared from a dry blend of stavudine, a spheronizing agent, a suitable diluent and a stabilizing amount of magnesium stearate. The magnesium stearate, in contrast to other similar pharmaceutical adjuncts, has been found to stabilize stavudine against degradation due to hydrolysis in the presence of the limited amount of water necessary for the extrusion-spheronization process. Also included in the scope of the invention are hard gelatin capsules containing, in addition to the stavudine beadlets, similar beadlets containing other therapeutic agents utilized to treat retroviral infections.