(19) World Intellectual Property Organization International Bureau





(43) International Publication Date 15 March 2001 (15.03.2001)

PCT

(10) International Publication Number WO 01/17547 A3

(51) International Patent Classification⁷: 35/30, A61P 25/28, C12N 5/06

A61K 38/18,

(21) International Application Number: PCT/US00/24555

(22) International Filing Date:

7 September 2000 (07.09.2000)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

60/153,121

8 September 1999 (08.09.1999) U.

- (71) Applicants: GENETICS INSTITUTE, INC. [US/US]; 87 Cambridge Park Drive, Cambridge, MA 02140 (US). TRUSTEES OF BOSTON UNIVERSITY [US/US]; 147 Bay State Road, Boston, MA 02215 (US).
- (72) Inventors: BLUSZTAJN, Jan, Krzysztof; 21
 Mount Vernon Street, Cambridge, MA 02140 (US).
 LOPEZ-COVIELLA, Ignacio; 2 Gem Avenue, Brighton,
 MA 02135 (US). KRAUSS, Raul; 20 Holden Street,
 Cambridge, MA 02138-2022 (US). THIES, R., Scott; 4
 Miles Circle, Andover, MA 01810 (US).

- (74) Agents: RYAN, M., Andrea; c/o Brady, Kay, E., American Home Products Corporation, Patent Law Dept., One Campus Drive, Parsippany, NJ 07054 et al. (US).
- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, 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), 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: 27 September 2001

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.

01/17547 A3

(54) Title: BMP-9 COMPOSITIONS AND METHODS FOR INDUCING DIFFERENTIATION OF CHOLINERGIC NEURONS

ional Application No.

INTERNATIONAL SEARCH REPORT PCT/US 00/24555 A. CLASSIFICATION OF SUBJECT MATTER IPC 7 A61K38/18 A61K35/30 A61P25/28 C12N5/06 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) A61K C12N IPC 7 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) BIOSIS, EPO-Internal, WPI Data, PAJ C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to claim No. Citation of document, with indication, where appropriate, of the relevant passages Category ° WO 99 01159 A (UNIV UTAH RES FOUND) 15,16 χ 14 January 1999 (1999-01-14) page 9, line 18 -page 11, line 27 page 22, line 1 -page 23, line 8 page 48, line 14 -page 49, line 29 page 53, line 32 -page 54, line 24 claims 1-3,7-12,16-21,25-28,32-43,46-59 -/--

Further documents are listed in the continuation of box C.	χ Patent family members are listed in annex.
A document defining the general state of the art which is not considered to be of particular relevance E earlier document but published on or after the international filing date L document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) O document referring to an oral disclosure, use, exhibition or other means P document published prior to the international filing date but later than the priority date claimed	 'T' later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention 'X' document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone 'Y' document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. '&' document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the international search report
28 February 2001	13/03/2001
Name and mailing address of the ISA	Authorized officer
European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040. Tx. 31 651 epo nl. Fax: (+31-70) 340-3016	Stein, A

2

INTERNATIONAL SEARCH REPORT

Int ional Application No PCT/US 00/24555

	Ation) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication where appropriate, of the relevant passages	Relevant to claim No.
Category °	Citation of document, with indication, where appropriate, or the relevant passages	helevalit to claim No.
A	KALYANI ANJALI ET AL: "Spinal cord neuronal precursors generate multiple neuronal phenotypes in culture." JOURNAL OF NEUROSCIENCE, vol. 18, no. 19, 1 October 1998 (1998-10-01), pages 7856-7868, XP002161574 ISSN: 0270-6474 abstract page 7863, right-hand column, line 11 - line 34 page 7865; figure 6 page 7866, right-hand column, line 34 -page 7867, left-hand column, line 3 page 7867, left-hand column, line 3 page 7867, left-hand column, line 32 - line 35	15,16
А	WO 99 37320 A (CREATIVE BIOMOLECULES INC) 29 July 1999 (1999-07-29)	1,3,4, 8-10, 13-15
	the whole document	10.10
А	US 5 756 457 A (TORIUMI DEAN M ET AL) 26 May 1998 (1998-05-26) the whole document	15,16
A	LOPEZ-COVIELLA I ET AL: "Bone morphogenetic protein-9 modulates acetylcholine content and choline acetyltransferase and vesicular acetylcholine transporter gene expression in the SN56 murine septal cell line." JOURNAL OF PHYSIOLOGY PARIS, vol. 92, no. 5-6, October 1998 (1998-10), pages 460-461, XP000982562 Xth International Symposium on Cholinergic Mechanisms, Part III; Paris, France; 1998 ISSN: 0928-4257 the whole document	1-16
Α	WO 95 33830 A (CELESTE ANTHONY J ;GENETICS INST (US); ROSEN VICKI A (US); THIES S) 14 December 1995 (1995-12-14) the whole document, especially page 8 lines 22-25	1-14
Ρ,Χ	LOPEZ-COVIELLA IGNACIO ET AL: "Induction and maintenance of the neuronal cholinergic phenotype in the central nervous system by BMP-9." SCIENCE (WASHINGTON D C), vol. 289, no. 5477, 2000, pages 313-316, XP002161575 ISSN: 0036-8075 the whole document	1-16
	-/	

2

INTERNATIONAL SEARCH REPORT

Int ional Application No PCT/US 00/24555

C.(Continu	ontinuation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
P,X	LOPEZ-COVIELLA I ET AL: "Bone morphogenetic proteins (BMPs) as cholinergic differentiating factors for basal forebrain neurons." SOCIETY FOR NEUROSCIENCE ABSTRACTS, vol. 25, no. 1-2, 1999, page 517 XPO00982558 29th Annual Meeting of the Society for Neuroscience, Part 1; Miami Beach, Florida, USA; October 23-28, 1999 ISSN: 0190-5295 the whole document	1-16

2

INTERNATIONAL SEARCH REPORT

Inti ional Application No PCT/US 00/24555

Patent document cited in search report		Publication date	ł	Patent family member(s)	Publication date
WO 9901159	А	14-01-1999	AU EP ZA	8382398 A 1011732 A 9805887 A	25-01-1999 28-06-2000 26-04-1999
WO 9937320	Α	29-07-1999	AU EP	2330999 A 1047443 A	09-08-1999 02-11-2000
US 5756 4 57	Α	26-05-1998	AU CA EP FI JP NO WO	677866 B 7868294 A 2169191 A 0716610 A 960809 A 9501932 T 960711 A 9505846 A	08-05-1997 21-03-1995 02-03-1995 19-06-1996 22-02-1996 25-02-1997 22-02-1996 02-03-1995
WO 9533830	Α	14-12-1995	AU AU EP JP KR US	709869 B 2817095 A 0764208 A 10503927 T 255417 B 6034061 A	09-09-1999 04-01-1996 26-03-1997 14-04-1998 01-05-2000 07-03-2000