

(19) World Intellectual Property Organization  
International Bureau



(43) International Publication Date  
30 November 2006 (30.11.2006)

PCT

(10) International Publication Number  
**WO 2006/127966 A3**

(51) International Patent Classification:

*G01N 33/53* (2006.01)    *A01N 63/00* (2006.01)  
*G01N 33/567* (2006.01)    *A01N 65/00* (2006.01)  
*A61K 38/00* (2006.01)    *C12N 5/06* (2006.01)  
*A61K 9/22* (2006.01)    *C12N 5/16* (2006.01)  
*A01N 37/18* (2006.01)    *A61M 1/00* (2006.01)

(21) International Application Number:

PCT/US2006/020371

(22) International Filing Date: 25 May 2006 (25.05.2006)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:

60/684,340    25 May 2005 (25.05.2005)    US

(71) Applicant (*for all designated States except US*): **THE JOHNS HOPKINS UNIVERSITY** [US/US]; 3400 N. Charles Street, Baltimore, MD 21218 (US).

(72) Inventors; and

(75) Inventors/Applicants (*for US only*): **SCHNAAR, Ronald, L.** [US/US]; 9094 Goldamber Garth, Columbia, MD 21045 (US). **YANG, Lynda, J., S.** [US/US]; 8112 Stone Ridge Drive, Frederick, MD 21702 (US). **SCHRAMM, Lawrence, P.** [US/US]; 5012 Boxhill Lane, Baltimore, MD 21210 (US).

(74) Agents: **CORLESS, Peter, F.** et al.; Edwards Angell Palmer & Dodge LLP, P.O. Box 55874, Boston, MA 02205 (US).

(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*

(88) Date of publication of the international search report:

1 March 2007

*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: COMPOSITIONS AND METHODS FOR ENHANCING AXON REGENERATION

(57) Abstract: As described below, the present invention generally features compositions and methods for the treatment of CNS disease or injury. In particular, the invention provides methods and compositions for enhancing axonal outgrowth in a subject. In one embodiment, the invention enhances the success of CNS restorative surgery.



WO 2006/127966 A3

# INTERNATIONAL SEARCH REPORT

International application No.

PCT/US06/20371

## A. CLASSIFICATION OF SUBJECT MATTER

IPC: G01N 33/53( 2006.01);33/567( 2006.01);A61K 38/00( 2006.01);9/22( 2006.01);A01N 37/18( 2006.01);63/00( 2006.01);65/00( 2006.01);C12N 5/06( 2006.01);5/16( 2006.01);A61M 1/00( 2006.01)

USPC: 435/7.2,326;514/2;530/350;424/93.1;604/890.1,123;977/906

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)

U.S. : 435/7.2,326;514/2;530/350;424/93.1;604/890.1,123;977/906

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 practicable, search terms used)

STN: Biosis, Biotechds, Embase, CAPLUS, Medline, USPATFULL, search term: outgrowth, sialidase, gangliosides, sialoglycoprotein, polysialic acid, GD1a, GT1b, chondroitinase, injury, neurodegenerative, inhibitor, regeneration.

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 97/01352 A1 (RESEARCH FOUNDATION OF CUNY, HUNTER COLLEGE) 16 January 1997, especially p. 15, line 6 to p21, line35.; p. 31-p. 36, examples5-8; p. 37 claims 1-17, p.41, claims 26-30.	1-4, 6-9, 12, 18-22, 39-44, 46-47 ----- 13-17, 23-38, 45, 48
Y	WO 03/074080 A1 (CAMBRIDGE UNIVERSITY TECHNICAL SERVICES LIMITED) 12 September 2003, especially p. 33, example 1,p.52 claims1-9; p. 57 claims 37-38.	1-48
Y	CHAU et al. Chondroitinase ABC enhances axonal regrowth through Schwann cell-seeded guidance channels after spinal cord injury. FASEB J. 2003, especially p. 2, 3rd paragraph.	13-17, 23-38, 45, 45-48



Further documents are listed in the continuation of Box C.



See patent family annex.

* Special categories of cited documents:		"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
"A"	document defining the general state of the art which is not considered to be of particular relevance	"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
"E"	earlier application or patent published on or after the international filing date	"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
"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)	"&"	document member of the same patent family
"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		

Date of the actual completion of the international search

14 September 2006 (14.09.2006)

Date of mailing of the international search report

01 NOV 2006

Name and mailing address of the ISA/US

Mail Stop PCT, Attn: ISA/US  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, Virginia 22313-1450

Facsimile No. (571) 273-3201

Authorized officer

Chang-Yu Wang

Telephone No. 571-272-1600

# INTERNATIONAL SEARCH REPORT

International application No.  
PCT/US06/20371

## C. (Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y,P	WO 2005/074655 A2 (EMORY UNIVERSITY) 18 August 2005, especially p.28, example 6 and example 7; pl 31, claims 1-24.	1-48
X	WARRINGTON et al. Neuron-Binding Monoclonal Antibodies Support Central Nervous System Neurite Extension. J. Neurophathol. Exp. Neurol. 2004, Vol. 63: 461-473, especially p. 464, 1st col. to 2nd col., 2nd paragraph; p. 468, 1st col. 1st paragraph.	1-9, 39-44 ----- 10-38, 45-48
Y	DEBELLARD et al. Myelin-Associated Glycoprotein Inhibits Axonal Regeneration from a Variety of Neurons via Interaction with a Sialoglycoprotein. Mol. Cell. Neurosci. 1996, Vol. 7: 89-101, especially p. 89, abstract; p. 98, 2nd paragraph to p. 99, 1st col., 2nd paragraph; p. 94 2nd col. to p. 95, 2nd col.	1-48
Y	ZUO et al. Degradation of Chondroitin Sulfate Proteoglycan Enhances the Neurite-Promoting Potential of Spinal Cord Tissue. Exp. Neurol. 1998, Vol. 154: 654-662, especially p. 654, abstract; p. 655, 1st col., 2nd paragraph.	13-17, 23-38, 45-47
X --- Y	TANG et al. Myelin-associated Glycoprotein Interacts with Neurons via a Sialic Acid Binding Site at ARG118 and a Distinct Neurite Inhibition Site. J. Cell Bio. 1997, Vol. 138: 1355-1366, especially p. 1355, abstract; p. 1357, 1st col., 5th paragraph to 2nd col. 4th paragraph; p. 1358, 2nd col.	1-9, 39-44. ----- 10-38, 45-48
Y	MCKEON et al. Exp. Injury-Induced Proteoglycans Inhibit the Potential for Laminin-Mediated Axon Growth on Astrocytic Scars. Neurol. 1995, Vol. 136: 32-43, especially p. 32, abstract; p. 33, 1st col. materials and methods.	13-17, 23-38, 45-47
A	MONNIER et al. The Polysialic Acid Moiety of the Neural Cell Adhesion Molecule is Involved in Intraretinal Guidance of Retinal Ganglion Cell Axons. Dev. Biol. 2001, Vol. 229: 1-14, especially p. 1, abstract.	1-48
Y	YANG et al. Gangliosides are neuronal ligands for myelin-associated glycoprotein. Proc. Natl. Acad. Sci. USA. 1996, Vol. 93: 814-818, especially p. 814, abstract; p. 815, 1st col.	1-48
A	PETRIDIS et al. Polysialic Acid Regulates Cell Contact-Dependent Neuronal Differentiation of Progenitor Cells from the Subventricular Zone. Dev. Dyn. 2004, 230: 675-684, p. 675 abstract.	1-48
Y,P	WO 2005/122734 A2 (THE RESEARCH FOUNDATION OF STATE UNIVERSITY OF NEW YORK) 29 December 2005, especially p.30-p32.	1-48