## (19) World Intellectual Property Organization

International Bureau



# T (BERGE BINGER) / FREIN BERGE (FELL III THE NICH BINGE BINGE RICH BERGER BERGER BERGER BERGER BERGER BERGER B

#### (43) International Publication Date 25 May 2001 (25.05.2001)

#### (10) International Publication Number WO 01/36638 A3

(51) International Patent Classification7: C12N 15/12, C07K 14/47, G01N 33/53, C12Q 1/68, A61K 38/17, 31/70, 39/395, A01K 67/027

Sheep Pasture Road, Madison, CT 06443 (US). VERNET, Corine [FR/US]; 4830 NW 43rd Street P#253, Gainesville, FL 32060 (US). FERNANDES, Elma [IN/US]; 77 Florence Road, #2B, Branford, CT 06405 (US).

- (21) International Application Number: PCT/US00/31543
- (22) International Filing Date:

17 November 2000 (17.11.2000)

(74) Agent: ELRIFI, Ivor, R.; Mintz, Levin, Cohn, Ferris, Glovsky and Popeo, P.C., One Financial Center, Boston, MA 02111 (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,

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

60/166,336	19 November 1999 (19.11.1999)	US
60/167,785	29 November 1999 (29.11.1999)	US
60/187,844	8 March 2000 (08.03.2000)	US
09/715,417	16 November 2000 (16.11.2000)	US

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, US, UZ, VN, YU, ZA, ZW.

(63) Related by continuation (CON) or continuation-in-part (CIP) to earlier applications:

US 60/166,336 (CIP) Filed on 19 November 1999 (19.11.1999) US 60/167,785 (CIP) Filed on 29 November 1999 (29.11.1999) US 60/187,844 (CIP) Filed on 8 March 2000 (08.03.2000) US Not furnished (CIP) Filed on 16 November 2000 (16.11.2000)

(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).

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

#### Published:

with international search report

(71) Applicant (for all designated States except US): CURA-

GEN CORPORATION [US/US]; 555 Long Wharf Drive, 11th floor, New Haven, CT 06511 (US).

(15) Information about Correction:

**Previous Correction:** 

see PCT Gazette No. 40/2001 of 4 October 2001, Section Π

10 May 2002

(72) Inventors; and

(75) Inventors/Applicants (for US only): SHIMKETS. Richard, A. [US/US]; 191 Leete Street, West Haven, CT 06516 (US). LICHENSTEIN, Henri [US/US]; 24

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: POLYPEPTIDES AND NUCLEIC ACIDS ENCODING SAME

(57) Abstract: The present invention provides novel isolated NOVX polynucleotides and polypeptides encoded by the NOVX polynucleotides. Also provided are the antibodies that immunospecifically bind to a NOVX polypeptide or any derivative, variant, mutant or fragment of the NOVX polypeptide, polynucleotide or antibody. The invention additionally provides methods in which the NOVX polypeptide, polynucleotide and antibody are utilized in the detection and treatment of a broad range of pathological states, as well as to other uses.



#### INTERNATIONAL SEARCH REPORT

International Application No Pc./US 00/31543

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12N15/12 C07K14/47 G01N33/53 C12Q1/68 A61K38/17 A61K39/395 A61K31/70 A01K67/027 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) IPC 7 C12N C07K G01N C12Q A61K 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) C. DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. Χ DATABASE EMBL [Online] 1-14,19 Entry/Acc.no. AI613267, 26 April 1999 (1999-04-26) STRAUSBERG, R.: "ty35d07.x1 NCI CGAP Ut2 Homo sapiens cDNA clone IMAGE: 2281069 3'. mRNA sequence." XP002174078 the whole document Α IKEGAWA SHIRO ET AL: "Structure and chromosomal assignment of the human S1-5 gene (FBNL) that is highly homologous to fibrillin" GENOMICS, ACADEMIC PRESS, SAN DIEGO, US. vol. 35, no. 3, 1996, pages 590-592. XP002155261 ISSN: 0888-7543 the whole document Further documents are listed in the continuation of box C. Patent family members are listed in 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 "A" document defining the general state of the art which is not considered to be of particular relevance invention "E" earlier document but published on or after the international filing date "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 "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) "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. "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 "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 27. 11. 01 7 August 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 ní, Fax: (+31-70) 340-3016 Smalt, R

### INTERNATIONAL SEARCH REPORT

International Application No
Pc./US 00/31543

		10.703	00/31543
C.(Continua Category °	ation) DOCUMENTS CONSIDERED TO BE RELEVANT  Citation of document, with indication, where appropriate, of the relevant passages		
oalegury *	onation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
A	WO 98 04688 A (BECHTOLD ROLF ;BIOPHARM GMBH (DE); POHL JENS (DE); UNSICKER KLAUS) 5 February 1998 (1998-02-05)		
4	WO 95 18823 A (BETH ISRAEL HOSPITAL) 13 July 1995 (1995-07-13)		
Ρ,χ	NAGASE T ET AL: "PREDICTION OF THE CODING SEQUENCES OF UNIDENTIFIED HUMAN GENES.17. THE COMPLETE SEQUENCES OF 100 NEW CDNA CLONES FROM BRAIN WHICH CODEFOR LARGE PROTEINS IN VITRO"  DNA RESEARCH, UNIVERSAL ACADEMY PRESS, JP, vol. 7, 2000, pages 143-150, XP000943428 ISSN: 1340-2838 * see KIAA1482 *		5-7, 10-14, 19,30,33
			i

### INTERNA I IONAL SEARCH REPORT

ernational application No. PCT/US 00/31543

Box i	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)	_
This Inte	rnational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:	
1. X	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:	
	Although claims 23-28, 42-43, and claim 22 in as far as it partains to in vivouse, are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.	)
	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:	
<u> </u>	Claims Nos.: pecause they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).	
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)	_
This Intern	national Searching Authority found multiple inventions in this international application, as follows:	
:	see additional sheet	
1. A	s all required additional search fees were timely paid by the applicant, this International Search Report covers all earchable claims.	
2. As	s all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment f any additional fee.	
3. As	s only some of the required additional search fees were timely paid by the applicant, this International Search Report overs only those claims for which fees were paid, specifically claims Nos.:	
	o required additional search fees were timely paid by the applicant. Consequently, this International Search Report is stricted to the invention first mentioned in the claims; it is covered by claims Nos.:  -43 all partially	
Remark on	Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.	

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1-43, all partially

NOV1 polypeptide and variants thereof as illustrated by seq.ID.2, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

2. Claims: 1-43, all partially

NOV2 polypeptide and variants thereof as illustrated by seq.ID.4, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

3. Claims: 1-43, all partially

NOV3 polypeptide and variants thereof as illustrated by seq.ID.6, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

4. Claims: 1-43, all partially

NOV4 polypeptide and variants thereof as illustrated by seq.ID.8, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich

binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 5. Claims: 1-43, all partially

NOV5 polypeptide and variants thereof as illustrated by seq.ID.10, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 6. Claims: 1-43, all partially

NOV6 polypeptide and variants thereof as illustrated by seq.ID.12, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 7. Claims: 1-43, all partially

NOV7 polypeptide and variants thereof as illustrated by seq.ID.14, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

## 8. Claims: 1-43, all partially

NOV8 polypeptide and variants thereof as illustrated by seq.ID.16, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide,

use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 9. Claims: 1-43, all partially

NOV9 polypeptide and variants thereof as illustrated by seq.ID.18, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 10. Claims: 1-43, all partially

NOV10 polypeptide and variants thereof as illustrated by seq.ID.20, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 11. Claims: 1-43, all partially

NOV11 polypeptide and variants thereof as illustrated by seq.ID.22, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 12. Claims: 1-43, all partially

NOV12 polypeptide and variants thereof as illustrated by seq.ID.24, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 13. Claims: 1-43, all partially

NOV13 polypeptide and variants thereof as illustrated by seq.ID.26, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 14. Claims: 1-43, all partially

NOV14 polypeptide and variants thereof as illustrated by seq.ID.28, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

#### 15. Claims: 1-43, all partially

NOV15 polypeptide and variants thereof as illustrated by seq.ID.30, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical

compositions of said polypeptide, nucleic acid, antibody or binding agent.

16. Claims: 1-43, all partially

NOV16 polypeptide and variants thereof as illustrated by seq.ID.32, nucleic acids encoding them and fragments thereof, vector comprising said nucleic acid, cell comprising said vector, antibody against said polypeptide, use of the antibody or the nucleic acid in a detection assay for said protein resp. encoding nucleic acid, method for diagnosing a disorder associated with altered levels of said polypeptide resp. nucleic acid, method for identifying an agent wich binds to said polypeptide, and pharmaceutical compositions of said polypeptide, nucleic acid, antibody or binding agent.

### INTERNATIONAL SEARCH REPORT

iformation on patent family members

International Application No Pt. /US 00/31543

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 9804688	A	05-02-1998	AU WO EP JP	4202897 A 9804688 A1 0922101 A1 2000516457 T	20-02-1998 05-02-1998 16-06-1999 12-12-2000
WO 9518823	A	13-07-1995	US WO US	5532167 A 9518823 A2 6004757 A	02-07-1996 13-07-1995 21-12-1999