

### **PCT**

# WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



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

(51) International Patent Classification 5: C12N 15/86, 5/10, 7/04 C12N 15/12, 15/62, 15/00 A01K 67/00

A3

(11) International Publication Number:

WO 92/05266

(43) International Publication Date:

2 April 1992 (02.04.92)

(21) International Application Number:

PCT/US91/06852

(22) International Filing Date:

20 September 1991 (20.09.91)

(30) Priority data:

586,603

21 September 1990 (21.09.90) US

(71) Applicant: VIAGENE, INC. [US/US]; 11075 Roselle Street, San Diego, CA 92121 (US).

(72) Inventors: JOLLY, Douglas, J.; 3050 Via Alicante Drive, La Jolla, CA 92037 (US). BARBER, Jack, R.; 11168 Carlotta Street, San Diego, CA 92129 (US). RESPESS, James, G.; 4966 Lamont Street, San Diego, CA 92109 (US). CHANG, Stephen, M., W.; 9838 Via Caceras, San Diego, CA 92129 (US). (74) Agents: MAKI, David, J. et al.; Seed and Berry, 6300 Columbia Center, Seattle, WA 98104-7092 (US).

(81) Designated States: AT (European patent), AU, BB, BE (European patent), BF (OAPI patent), BG, BJ (OAPI patent), BR, CA, CF (OAPI patent), CG (OAPI patent), CH (European patent), CI (OAPI patent), CM (OAPI patent), CS, DE (European patent), DK (European patent), ES (European patent), FI, FR (European patent), GA (OAPI patent), GB (European patent), GN (OAPI patent), GR (European patent), HU, IT (European patent), JP, KP, KR, LK, LU (European patent), MC, MG, ML (OAPI patent), MN, MR (OAPI patent), MW, NL (European patent), NO, PL, RO, SD, SE (European patent), SN (OAPI patent), TG (OAPI patent), 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 the receipt of amendments.

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

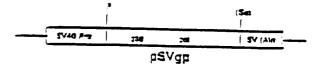
23 July 1992 (23.07.92)

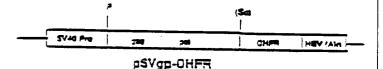
#### (54) Title: PACKAGING CELLS

#### (57) Abstract

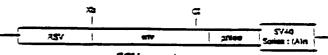
The invention described herein allows the production of recombinant retroviruses (retroviral vector particles) from producer cells which are safer and of higher titre than normal. In addition, methods are provided for making helper cells which, when a recombinant retrovirus genome is introduced to make a producer line, produce particles that are targeted toward particular cell types. Methods are also provided for making recombinant retrovirus systems adapted to infect a particular cell type, such as a tumor, by binding the retrovirus or recombinant retrovirus in the particular cell type. Methods are also provided for producing recombinant retroviruses which integrate in a specific small number of places in the host genome, and for producing recombinant retroviruses from transgenic animals.

# Plasmids Designed to Increase Viral Frotein Production









pRSVenv-gled

### FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AT	Austria	FI	Finland	ML	Mali
AU	Australia	FR	France	MN	Mongolia
88	Barbados	GA	Gubon	MR	Mauritania
BE	Belgium	CB	United Kingdom	MW	Malawi
BF	Burkina Faso	GN	Guinea	NI.	Netherlands
BG	Bulgaria	GR	Greece	NO	Norway
BJ	Benin	HU	Hungary	PL	Poland
BR	Brazil	ΙE	Ireland	RO	Romania
CA	Canada	ıT	Italy	RU	Russian Federation
CF	Central African Republic	jР	Japan	SD	Sudan
CG	Congo	KP	Democratic People's Republic	SE	Sweden
СН	Switzerland		of Korca	SN	Senegal
CI	Côte d'Ivoire	KR	Republic of Korea	SU	Soviet Union
СМ	Cameroon	LI	Liechtenstein	TD	Chad
CS	Czechoslovakia	LK	Sri Lanka	TG	Togo
DE	Germany	LU	Luxembourg	US	United States of America
ÐK	Denmark	MC	Monaco		
ES	Spain	MG	Madagascar		

## INTERNATIONAL SEARCH REPORT

International Application No

PCT/US 91/06852

	ECT MATTER (if several classification sy		
Int.Cl.5	t Classification (IPC) or to both National Cla C 12 N 15/86 C 12 C 12 N 15/62 C 12	2N 5/10 C 12N 7/	· · · · · · · · · · · · · · · · · · ·
II. FIELDS SEARCHED			
	Minimum Docume	ntation Searched <sup>7</sup>	
Classification System		Classification Symbols	
Int.Cl.5	C 12 N		
	Documentation Searched other to the Extent that such Documents a	than Minimum Documentation are Included in the Fields Searched <sup>8</sup>	
III. DOCUMENTS CONSIDER	ED TO BE RELEVANT <sup>9</sup>		
Category ° Citation of I	Document, 11 with indication, where appropris	nte, of the relevant passages 12	Relevant to Claim No.13
1991, P,Y * See X WO,A, Novem Y see t A Nucle 1990, mamma vecto a com	9102805 (VIAGENE, INC.) see page 12; page 13, 1 page 84, line 26 - page 8808454 (STATE OF OREGO ber 1988, see especially he whole document  ic Acids Research, vol. J.P. MORGENSTERN et al. lian gene tranfer: high rs with multiple drug se plementary helper-free pages 1587-3596, see the who	lines 1-4; example 8 2 86, line 7 *  ON) 3 7 table I  18, no. 12, 25 June 18. "Advanced 18. titre retroviral 20 election markers and 20 packaging cell line",	1 13-23 20,22 13-23 1
"E" earlier document but pu filling date "L" document which may th which is cited to establi citation or other special "O" document referring to a other means	general state of the art which is not icular relevance bilished on or after the international row doubts on priority claim(s) or sh the publication date of another reason (as specified) an oral disclosure, use, exhibition or or to the international filling date but late claimed	"T" later document published after the intern or priority date and not in conflict with a cited to understand the principle or theor invention  "X" document of particular relevance; the cla cannot be considered novel or cannot be involve an inventive step  "Y" document of particular relevance; the cla cannot be considered to involve an inventive an inventive an inventive an inventive document is combined with one or more ments, such combination being obvious a in the art.  "A" document member of the same patent far.  Date of Mailing of this international Sea	he application but y underlying the imed invention considered to imed invention tive step when the other such docus- o a person skilled mily
	PEAN PATENT OFFICE	MASHI MORNIEUSEN	!

International Application No Page 2 PCT/US 91/06852

	IS CONSIDERED TO BE RELEVANT (CONTINUED FROM THE SECOND SHEET)	
Category °	Citation of Document, with indication, where appropriate, of the relevant passages	Relevant to Claim No.
(	EP,A,0386882 (DANA FARBER CANCER	20-21
	INSTITUTE) 12 September 1990, see the whole	
	document	
		1 00 01
(	Journal of Virology, vol. 64, no. 3, March 1990,	1,20-21
	FL. COSSET et al.: "A new avian leukosis	
	virus-based packaging cell line that uses two	
	separate transcomplementing helper genomes",	
	pages 1070-1078, see the whole document,	
	specially page 1076, column 2	
Ì		
ļ		

	International A	ication No. PCT/ US91 /06852
FURTHER INFORMATION CONTINUED FROM THE SECOND S	HEET	
V. OBSERVATION WHERE CERTAIN CLAIMS WERE FOU	ND UNSEARCHABLE 1	
This International search report has not been established in respect of certain	claims under Article 17(2)(a) fo	or the following reasons:
		er not required to be searched by this
Authority, namely:	•	
with the prescribed requirements to such an extent that no meaningful	international search can be ca	International application that do not comply irried out, specifically:
A NO CONTRACTOR WATER INVENTION OF INVENTION	CKING 2	
VI.X. OBSERVATIONS WHERE UNITY OF INVENTION IS LAC		
This International Searching Authority found multiple Inventions in this Internation 1. Claims 1,13-23; 2. Claims 2,3 3. Claims 4-7,31; 4. Claim 8 5. Claims 9-12; 6. Claims 24-30	kional application as follows:	
As all required additional search fees were timely paid by the applicant of the International application.	t, this internetional search rep	ort covers all searchable claims
As only some of the required additional search fees were timely paid those claims of the International application for which fees were paid,	y the applicant, this internatio specifically claims:	nal search report covers only
No required additional search fees were timely paid by the applicant. On the invention first mentioned in the claims; it is covered by claim num Claims 1, 13-23	Consequently, this international bers:	I search report is restricted to
As all searchable claims could be searched without effort justifying an invite payment of any additional fee.  Remark on Protest	additional fee, the Internation	al Searching Authority did not
The additional search fees were accompanied by applicant's protest.  No protest accompanied the payment of additional search fees.		

### ANNEX TO THE INTERNATIONAL SEARCH REPORT ON INTERNATIONAL PATENT APPLICATION NO.

US 9106852

SA 52929

This annex lists the patent family members relating to the patent documents cited in the above-mentioned international search report. The members are as contained in the European Patent Office EDP file on 12/06/92

The European Patent Office is in no way liable for these particulars which are merely given for the purpose of information.

Patent document cited in search report	Publication date	Patent family member(s)		Publication date
WO-A- 9102805	07-03-91	AU-A- EP-A-	6185390 0487587	03-04-91 03-06-92
WO-A- 8808454	03-11-88	AU-A-	1782288	02-12-88
EP-A- 0386882	12-09-90	CA-A- JP-A-	2009403 3061492	06-08-90 18-03-91