(19) World Intellectual Property Organization

International Bureau





(43) International Publication Date 21 August 2008 (21.08.2008)

(10) International Publication Number WO 2008/098693 A3

(51) International Patent Classification:

C07K 1/06 (2006.01) C07K 7/04 (2006.01) C07K 1/10 (2006.01) C07K 7/06 (2006.01) C07K 5/06 (2006.01) A61K 47/48 (2006.01)

(21) International Application Number:

PCT/EP2008/000846

(22) International Filing Date: 2 February 2008 (02.02.2008)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:

07003302.2 16 February 2007 (16.02.2007) EP 07005646.0 20 March 2007 (20.03.2007)

(71) Applicant (for all designated States except US): BAYER HEALTHCARE AG [DE/DE]; 51368 Leverkusen (DE).

(72) Inventors; and

- (75) Inventors/Applicants (for US only): KRUGER, Joachim [DE/DE]; Zeppenheimer Weg 71, 40489 Dusseldorf (DE). MILITZER, Hans-Christian [DE/DE]; Waldweg 2, 51519 Odenthal (DE). **SIEGEL, Konrad** [DE/DE]; $G\delta r$ resstr. 13, 40597 Dusseldorf (DE). MINUTH, Torsten [DE/DE]; Wackenbergstrasse 58a, 13156 Berlin (DE).
- (74) Common Representative: BAYER HEALTHCARE AG; Law and Patents, Patents and licensing, 51368 Leverkusen (DE).

- (81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, 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, HR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL, NO, 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
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments
- with sequence listing part of description published sep arately in electronic form and available upon request from the International Bureau
- (88) Date of publication of the international search report:

20 November 2008

(54) Title: CONVERGENT SYNTHESIS OF GLP-I

(57) Abstract: The present invention relates to a process for the manufacture of the polypeptide GLP-I (SEQ ID NO 1) and its PEGylated derivative, bei fragment condensation (convergent synthesis).

INTERNATIONAL SEARCH REPORT

International application No PCT/EP2008/000846

Δ	CLASSIRCATION	OF SUBJECT	MATTER
м.	CLASSINGATION	OF SUBJECT	WAITER

C. DOCUMENTS CONSIDERED TO BE RELEVANT

A61K47/48

INV. C07K1/06

C07K1/10

C07K5/06

C07K7/04

C07K7/06

Relevant to claim No.

1-19

According to International Patent Classification (IPC) orto both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

C07K

Category*

Y

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)

EPO-Internal , WPI Data, EMBASE, BIOSIS, Sequence Search

Citation of document, with indication, where appropriate, of the relevant passages

WO 03/040309 A (BAYER AG [US]; PAN CLARK [US]; WHELAN JAMES [US]; CLAIRMONT KEVIN B

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) 10° 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Cited to understand the pinvention of tive ant winch is not ant within in the art winch is considered to cannot be considered no involve an inventive step document of particular relevance to exhibition or obcument is combined with ments, such combination in the art. "8.1 document member of the document member of the international search Date of mailing of the international officer Authorized officer Ravrak Sin	
the whole document WO 01/04156 A (ZEALAND PHARMACEUTICALS AS [DK]; LARSEN BJARNE DUE [DK]; MIKKELSEN JEN) 18 January 2001 (2001-01-18) page 22, line 29 - line 33 WO 2006/087354 A (NOVO NORDISK AS [DK]; BEHRENS CARSTEN [DK]; LAU JESPER [DK]; KODRA JAN) 24 August 2006 (2006-08-24) The 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 (almost be considered to the considered to the considered to other means of the means (as specified) Codocument referring to an oral disclosure, use, exhibition or other means (as specified) Date of the actual completion of the international filing date but later than the priority date claimed Date of the actual completion of the international search September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk, Tel. (+31-70) 340-2040, Tx. 31 651 epo ni. Tater document family and or priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) To 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) To document therefore the considered to document is combined in in the an. Standard Televant Televa	
WO 01/04156 A (ZEALAND PHARMACEUTICALS AS [DK]; LARSEN BJARNE DUE [DK]; MIKKELSEN JEN) 18 January 2001 (2001-01-18) page 22, line 29 - line 33 WO 2006/087354 A (NOVO NORDISK AS [DK]; BEHRENS CARSTEN [DK]; LAU JESPER [DK]; KODRA JAN) 24 August 2006 (2006-08-24) D Further documents are listed in the continuation of Box C. Special categories of cited documents: 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 10 document published prior to the international filing date of another citation or other special reason (as specified) 10 document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (431-70) 340-2040, Tx. 31 651 epo nl, Rayrak Si.	
[DK]; LARSEN BJARNE DUE [DK]; MIKKELSEN JEN) 18 January 2001 (2001-01-18) page 22, line 29 - line 33 WO 2006/087354 A (NOVO NORDISK AS [DK]; BEHRENS CARSTEN [DK]; LAU JESPER [DK]; KODRA JAN) 24 August 2006 (2006-08-24) **Special categories of cited documents:* A document defining the general state of the art which is not considered to be of particular relevance Te earlier document but published on or after the international filing date Tu-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) To document referring to an oral disclosure, use, exhibition or other means The document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (4-31-70) 340-2040, Tx. 31 651 epo nl. Ravrak Si.	
[DK]; LARSEN BJARNE DUE [DK]; MIKKELSEN JEN) 18 January 2001 (2001-01-18) page 22, line 29 - line 33 WO 2006/087354 A (NOVO NORDISK AS [DK]; BEHRENS CARSTEN [DK]; LAU JESPER [DK]; KODRA JAN) 24 August 2006 (2006-08-24) **Special categories of cited documents:* A document defining the general state of the art which is not considered to be of particular relevance Te earlier document but published on or after the international filing date Tu-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) To document referring to an oral disclosure, use, exhibition or other means The document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (4-31-70) 340-2040, Tx. 31 651 epo nl. Ravrak Si.	
A WO 2006/087354 A (NOVO NORDISK AS [DK]; BEHRENS CARSTEN [DK]; LAU JESPER [DK]; KODRA JAN) 24 August 2006 (2006-08-24) A document defining the general state of the art which is not considered to be of particular relevance earlier document but published on or after the international filing date which is cited to establish the publication date of another citation or other special reason (as specified) 10' document referring to an oral disclosure, use, exhibition or other means 19' document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl. Pavrak Si	1-19
MO 2006/087354 A (NOVO NORDISK AS [DK]; BEHRENS CARSTEN [DK]; LAU JESPER [DK]; KODRA JAN) 24 August 2006 (2006-08-24) To special categories of cited documents: A document defining the general state of the art which is not considered to be of particular relevance E artier 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) 10' 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Fel. (+317-70) 340-2040, Tx. 31 651 epo nl, Rawrak Si.	
WO 2006/087354 A (NOVO NORDISK AS [DK]; BEHRENS CARSTEN [DK]; LAU JESPER [DK]; KODRA JAN) 24 August 2006 (2006-08-24) Special categories of cited documents: A document defining the general state of the art which is not considered to be of particular relevance if E earlier document but published on or after the international filling 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 filling date but later than the priority date claimed Date of the actual completion of the international search Date of mealth actual completion of the international search September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,	
BEHRENS CARSTEN [DK]; LAU JESPER [DK]; KODRA JAN) 24 August 2006 (2006-08-24) The special categories of cited documents: 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 filling 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) 10' document referring to an oral disclosure, use, exhibition or other means P' document published prior to the international filling date but later than the priority date claimed Date of the actual completion of the international search 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (431-70) 340-2040, Tx. 31 651 epo nl, Tater document published or priority date and not in cited to understand the prio	
BEHRENS CARSTEN [DK]; LAU JESPER [DK]; KODRA JAN) 24 August 2006 (2006-08-24) The special categories of cited documents: 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 filling 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) 10' document referring to an oral disclosure, use, exhibition or other means P' document published prior to the international filling date but later than the priority date claimed Date of the actual completion of the international search 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (431-70) 340-2040, Tx. 31 651 epo nl, Tater document published or priority date and not in cited to understand the prio	1
Further documents are listed in the continuation of Box C. Special categories of cited documents: 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) 10' 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, See patent family and Take document published or priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date on priority claim(s) or which is cited to establish the publication date of another claim(s) or document of particular relevance and the priority date considered to document is considered to document is combined to ments, such combination in the art. *** document of particular relevance and the priority date claimed and not in cited to understand the priority date considered no involve an inventive step 'Y' document of particular relevance and the priority date considered to document is considered to document is considered to document is considered to document of particular relevance and the priority date claimed and not in cited to understand the priority date claimed and not in cited to understand the priority date claimed and not in cited to understand the priority date claimed and not in cited to understand the priority date claimed and not in cited to understand the priority date claimed and priority date claimed and not in cited to understand the priorit	
Further documents are listed in the continuation of Box C. X See patent family and Special categories of cited documents: 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 C 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) C 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 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, T later document published or priority date and not in cited to understand the priority date considered to considered to cannot be considered to document of particular releannt be considered to document is combined when the priority date claimed	
Special categories of cited documents: 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) 10° 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, T' later document published or priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority document of particular relevance "X" document of particular relevance "Y' document of particular relevance "A document of particular relevance "Y' document of particu	
Special categories of cited documents: 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) 10° 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, T' later document published or priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority document of particular relevance "X" document of particular relevance "Y' document of particular relevance "A document of particular relevance "Y' document of particu	
Special categories of cited documents: 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) 10° 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, T' later document published or priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority document of particular relevance "X" document of particular relevance "Y' document of particular relevance "A document of particular relevance "Y' document of particu	
Special categories of cited documents: 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) 10° 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, T' later document published or priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority document of particular relevance "X" document of particular relevance "Y' document of particular relevance "A document of particular relevance "Y' document of particu	
Special categories of cited documents: 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) 10° 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, T' later document published or priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority document of particular relevance "X" document of particular relevance "Y' document of particular relevance "A document of particular relevance "Y' document of particu	
Special categories of cited documents: 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) 10° 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, T' later document published or priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority date and not in cited to understand the priority document of particular relevance "X" document of particular relevance "Y' document of particular relevance "A document of particular relevance "Y' document of particu	
'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) 10' 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Talter document published or or priority date and not in cited to understand the priority date on or involve an inventive step document of particular relevanto be considered no involve an inventive step document is combined we ments, such combination in the art. "&1 document of particular relevanto be considered no involve an inventive step document is combined we ments, such combined we ments, such combination in the art. "&1 document of particular relevanto be considered no involve an inventive step document is combined we ments, such combined we ments, such combined we ments, such combined we ments, such combined to document is combined to documen	nex.
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) 10' 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Por document defining the general state of the art which is not cited to understand the picted to indected to understand the picted	
"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) 10° 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Ravrak Sin	after the international filing date conflict with the application but rinciple or theory underlying the
"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) 10° document referring to an oral disclosure, use, exhibition or other means 1P' document published prior to the international filling date but later than the priority date claimed 1September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Rannot be considered no involve an involve and involv	evance the claimed invention
which is cited to establish the publication date of another citation or other special reason (as specified) 10' document referring to an oral disclosure, use, exhibition or other means 1P' document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Authorized officer Py' document of particular relicannot be considered to document is combined we ments, such combination in the art. "&1 document member of the document member of the means of the lateral document member of the means of the international search Date of mailing of the international search Authorized officer Rayrak Sin	vel or cannot be considered to
cannot be considered to 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 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Cannot be considered to document is combined to document is combined we ments, such combination in the art. "&1 document member of the Date of mailing of the international search Authorized officer Rayrak Si	evance; the claimed invention
other means other means ments, such combination in the art. "&1 document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 5 September 2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Rayrak Sin	involve an inventive step when the vith one or more other such docu-
Date of the actual completion of the international search Date of the actual completion of the international search Date of mailing of the international search Date of mailing of the international search Date of mailing of the international search 08/10/2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Rayrak Sin	being obvious to a person skilled
5 September 2008 08/10/2008 Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Rayrak Si	same patent family .
Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Rayrak Si	rnational search report
Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Rayrak Si	
European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,	
NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Rayrak Si i	
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Rayrak Si	
	nasi
Fax: (+31-70) 340-3016	1401

International application No. PCT/EP2008/000846

INTERNATIONAL SEARCH REPORT

Box No. II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)							
This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:							
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:							
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:							
3. Lack Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).							
Box No. III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)							
This International Searching Authority found multiple inventions in this international application, as follows:							
see additi onal sheet							
As all required additional search fees were timely paid by the applicant, this international search report covers allsearchable claims.							
2. As all searchable claims could be searched without effort justifying an additional fees, this Authority did not invite payment of additional fees.							
3. As only some of the required additional search fees were timely paid by the applicant, this international search reportcovers only those claims for which fees were paid, specifically claims Nos.:							
4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-16, 18, 19							
Remark on Protest							
I No protest accompanied the payment of additional search fees.							

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

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

1. claims: 1-16,18,19

A process for the convergent synthesis of GLPl (SEQ ID NO 1) and the intermediate products.

2. claim: 17

A process for the preparation of PEGyI ated derivative of GLPl (SEQ ID NO 1). $\dot{}$

INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No
PCT/EP2008/000846

Patent document cited in search report		Publication date		Patent family member(s)		Publication date
wo 03040309	A	15-05-2003	BR	0213132	A	19-10-2004
		.0 00 =000	CA	2462462	Al	15-05-2003
			CN	1564828	Α	12-01-2005
			EP	1436330	A2	14-07-2004
			HK	1073117	Al	29-02-2008
			HR	20040327	A2	30-11-2007
			JР	2005508639	Т	07-04-2005
			KR	20050034580	Α	14-04-2005
			MA	26228	Al	01-07-2004
			MX	PA04002140	Α	07-03-2005
			NO	20041792	Α	02-07-2004
			NZ	532685	Α	24-02-2006
			UY	27469	ΑI	30-04-2003
			ZA	200403322	Α	03-05-2005
wo 0104156	Α	18-01-2001	AT	242267	T	15-06-2003
			ΑU	781338	B2	19-05-2005
			ΑU	5966000	Α	30-01-2001
			CA	2378431	Al	18-01-2001
			CN	1376166	Α	23-10-2002
			DE	60003182	DI	10-07-2003
			DE	60003182	12	06-05-2004
			DK	1196444	T3	29-09-2003
			EP	1196444	ΑI	17-04-2002
			EP	1329458	A2	23-07-2003
			EP	1076066	ΑI	14-02-2001
			ES	2200892	T3	16-03-2004
			JΡ	2003505347	Τ	12-02-2003
			JΡ	2007001987	Α	11-01-2007
			NZ	517012	Α	28-02-2003
			PT	1196444	T	31-10-2003
			US	2007111940	Al	17-05-2007
wo 2006087354	Α	24-08-2006	AU	2006215566	Al	24-08-2006
•			CA	2601745	ΑI	24-08-2006
			CN	101123992	Α	13-02-2008
			EP	1853321	A2	14-11-2007
			JP	2008530178	T	07-08-2008
			KR	20070108180	Α	08-11-2007