



(43) International Publication Date
5 December 2013 (05.12.2013)

(10) International Publication Number
WO 2013/181597 A3

- (51) **International Patent Classification:**
A61K 47/48 (2006.01) *A61K 51/10* (2006.01)
- (21) **International Application Number:**
PCT/US2013/043708
- (22) **International Filing Date:**
31 May 2013 (31.05.2013)
- (25) **Filing Language:** English
- (26) **Publication Language:** English
- (30) **Priority Data:**
61/653,823 31 May 2012 (31.05.2012) US
- (71) **Applicant:** SYNCHM, INC. [US/US]; 1400 Chase Avenue, Elk Grove Village, IL 60007 (US).
- (72) **Inventors:** XU, Ze-Qi; 6609 Chick Evans Lane, Woodbridge, IL 60517 (US). WANG, Qianli; 6835 N Dowagiac Avenue, Chicago, IL 60646 (US). JOSEPH, Augustine Rudolph; 2273 Webster Lane, Des Plaines, IL 60018 (US). FANG, Zhiqiang; 4 Charleston Court, Hawthorn Woods, IL 60047 (US).
- (74) **Agent:** LIBBY, Jelena Janjic; McDonnell Boehnen Hulbert & Berghoff LLP, 300 South Wacker Drive, Suite 3200, Chicago, IL 60606 (US).
- (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, BN, BR, BW, BY, BZ, CA, CH, CL, 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, 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, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, 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, LR, LS, MW, MZ, NA, RW, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MK, MT, NL, NO, PL, PT, RO, RS, SE, SI, SK, SM, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, KM, ML, MR, NE, SN, TD, TG).

Published:

- *with international search report (Art. 21(3))*
- *before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments (Rule 48.2(h))*

- (88) **Date of publication of the international search report:**
23 January 2014

(54) **Title:** THIOL-ENE CLICK CHEMISTRY FOR DRUG CONJUGATES

(57) **Abstract:** The present invention relates to linker molecules that readily conjugate cellular recognition ligand at one end and drug payload at the other, and are useful in treating or preventing cancer, an autoimmune disease, an inflammatory condition, a central nervous system disorder or an infection. The Invention includes linker molecules, Linker - Drug compounds, and Ligand - Linker - Drug conjugates.



WO 2013/181597 A3

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US13/43708

A. CLASSIFICATION OF SUBJECT MATTER

IPC(8) - A61K 39/40 (2013.01)

USPC - 424/178.1

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(8): A61K 39/40, 00; C07K 16/00; C08B 3/16 (2013.01)

USPC: 424/178.1, 179.1, 181.1, 184.1; 530/391.7, 391.9, 351; 536/64, 63

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)

MicroPatent (US-G, US-A, EP-A, EP-B, WO, JP-bib, DE-C,B, DE-A, DE-T, DE-U, GB-A, FR-A); PROQuest; Google; Google Scholar; Scirus; PubMed; PubChem; ligand-linker-drug conjugate; antibody drug conjugate; thiol-ene click chemistry; alkene; olefin; thiol; thioether; radioisotope; doxorubicin; auristatin

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	US 2003/0130189 A1 (SETER, PD et al.) 10 July 2003; abstract; paragraphs [0013], [0047], [0048], [0082], [0085], [0089], [0096]-[0098], [0100]-[0103], [0113]-[0117], [0121]-[0125], [0146], [0181]	1-3, 4/1-3, 6-9, 10/6-9, 11/6-9
Y	WO 2011/090428 A1 (CÓRDOVA, A et al.) 28 July 2011; abstract; scheme 3; paragraphs 2-3, page 1; paragraphs 1-4, page 2; paragraphs 1-5, page 5	1-3, 4/1-3, 6-9, 10/6-9, 11/6-9
Y	US 2011/0305631 A1 (GOVINDAN, SV et al.) 15 December 2011; abstract; paragraphs [0162], [0166]-[0169] and [0180]-[0182]	3, 4/3

☐ 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 application or patent 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

08 November 2011 (08.11.2013)

Date of mailing of the international search report

20 NOV 2013

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:

Shane Thomas

PCT Helpdesk: 571-272-4300
PCT OSP: 571-272-7774

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US13/43708

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:
2. ☐ 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. ☒ Claims Nos.: 5
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:

1. ☐ As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying 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 report covers 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.:

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee.
- ☐ The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation.
- ☐ No protest accompanied the payment of additional search fees.