



(51) International Patent Classification:

C12N 5/071 (2010.01) A61P 1/16 (2006.01)
A61K 35/12 (2015.01) A61P 9/04 (2006.01)
A61P 25/16 (2006.01)

(21) International Application Number:

PCT/SG2021/050412

(22) International Filing Date:

15 July 2021 (15.07.2021)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

63/054,206 20 July 2020 (20.07.2020) US

(71) Applicants: **CELLRESEARCH CORPORATION PTE LTD** [SG/SG]; 137 Market Street, #08-02, Grace Global Raffles Building, Singapore 048943 (SG). **SINGAPORE HEALTH SERVICES PTE LTD** [SG/SG]; Third Hospital Avenue, #03-03 Bowyer Block, Singapore 168753 (SG).

(72) Inventors: **CHAI, Chou**; 9H Surin Avenue, Singapore 534062 (SG). **LIM, Kah Leong**; 32, Braddell Road, #04-23 E-Maison, Singapore 359967 (SG). **PHAN, Toan Thang**;

137 Market Street, #08-02, Grace Global Raffles Building, Singapore 048943 (SG).

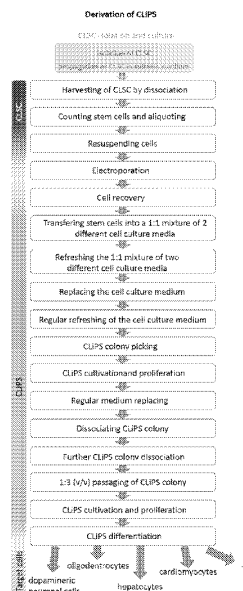
(74) Agent: **MCLAUGHLIN, Michael, Gerard et al.**; 112 Robinson Road, #14-041, Singapore 068902 (SG).

(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, DJ, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IR, IS, IT, JO, JP, KE, KG, KH, KN, KP, KR, KW, KZ, LA, LC, LK, LR, LS, 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, SA, SC, SD, SE, SG, SK, SL, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, WS, 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, ST, 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,

(54) Title: A METHOD OF GENERATING AN INDUCED PLURIPOTENT STEM CELL, AN INDUCED PLURIPOTENT STEM CELL AND METHODS OF USING THE INDUCED PLURIPOTENT STEM CELL

Figure 1:



(57) Abstract: The invention relates to a method of generating an induced pluripotent stem cell. The method of the disclosure comprises expressing exogenous nucleic acids encoding the proteins *OCT3/4*, *SOX2*, *KLF4*, *LIN28* and *L-MYC* and the *p53-shRNA* in a stem cell of the amniotic membrane of the umbilical cord under conditions suitable to reprogram the stem cell, thereby generating the induced pluripotent stem cell. The present invention also refer to an induced pluripotent stem cell population obtainable by the method and an induced pluripotent stem cell population obtained by the method. Further, a pharmaceutical composition comprising the induced pluripotent stem cell of the present invention is concerned. The present invention also relates to a method of differentiating the induced pluripotent stem cell of this invention. In addition, a pharmaceutical composition comprising a differentiated induced pluripotent stem cell obtained by the method is also concerned. Further, the present invention concerns a method of treating a congenital or acquired degenerative disorder in a subject, comprising administering to a subject a target cell differentiated from pluripotent stem cell.

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))*
- *with sequence listing part of description (Rule 5.2(a))*
- *in black and white; the international application as filed
contained color or greyscale and is available for download
from PATENTSCOPE*