

US010918695B2

(12) United States Patent

Sharma et al.

(54) USE OF JACK BEAN LECTIN FOR INCREASING THE ABUNDANCE OF HEMATOPOIETIC STEM CELLS AND PROGENITOR CELLS IN BONE MARROW AND/OR EPIDERMAL STEM CELLS IN SKIN IN VIVO

(71) Applicant: THE SECRETARY, DEPARTMENT OF ATOMIC ENERGY, Mumbai (IN)

(72) Inventors: Deepak Sharma, Mumbai (IN);
Santosh Kumar Sandur, Mumbai (IN);
Maikho Thoh, Mumbai (IN);
Raghavendra Shridhar Patwardhan,
Mumbai (IN); Dharmendra K.
Maurya, Mumbai (IN); Rahul
Checker, Mumbai (IN); Vikram P.
Gota, Navi Mumbai (IN); Jayakumar

Gota, Navi Mumbai (IN); Jayakum Sundarraj, Mumbai (IN); Haldhar Dev Sarma, Mumbai (IN); Subrata Chattopadhyay, Mumbai (IN)

Chattopadhyay, Mumbai (IN)

(73) Assignee: THE SECRETARY, DEPARTMENT OF ATOMIC ENERGY, Maharastra

(IN)

(*) Notice: Subject to any disclaimer, the term of this

patent is extended or adjusted under 35

U.S.C. 154(b) by 31 days.

(21) Appl. No.: 16/239,280

(22) Filed: Jan. 3, 2019

(65) **Prior Publication Data**

US 2020/0215152 A1 Jul. 9, 2020

(51) Int. Cl.

A61K 36/48 (2006.01)

A61K 38/16 (2006.01)

A61K 9/00 (2006.01)

(52) U.S. Cl.

(58) Field of Classification Search

CPC A61K 38/00; A61K 38/168; A61K 36/48; A61K 9/0019; C07K 14/415; C07K 14/42 See application file for complete search history.

(56) References Cited

FOREIGN PATENT DOCUMENTS

CA 2258503 12/1997

OTHER PUBLICATIONS

Fitzgerald et al. Comparison of the effects of concanavalin-A and epidermal growth factor on epithelial cell proliferation in the rat intestine. Aliment Pharmacol Ther 15: 1077-1084, 2001.* Lin et al. Concanavalin A (NSC-143504): Its action on experimental tumor cells and possible use in cancer chemotherapy. Cancer Chemotherapy Reports 59(2): 319-326, 1975.*

(10) Patent No.: US 10,918,695 B2

(45) **Date of Patent:** Feb. 16, 2021

Blasco E et al., "Proliferative response of human CD4' T lymphocytes stimulated by the lectin jacalin" European Journal of Immunology; Jul. 1995; vol. 25(7): pp. 2010-2018).

Boitano AE et al., "Aryl hydrocarbon receptor antagonists promote the expansionof human hematopoietic stem cell" Science Magazine, Sep. 10, 2010; vol. 329(5997):pp. 345-1348.

Bonomo et al., "A T Cell View of the Bone Marrow" May 2016; Frontiers in Immunology, 7: 184.

Brissot E et al., "Factors predicting allogeneic PBSCs yield after G-CSF treatment in healthy donors" Bone Marrow Transplant. Nov. 2009; vol. 44(9):pp. 613-615.

Broxmeyer HE et al., "Th1 cells regulate hematopoietic progenitor cell homeostasis by production of oncostatin M;" Immunity. 2002;16(6):815-25.

Greaves MF et al., "Activation of Human T and B Lymphocytes by Polyclonal Mitogens;" Nature Publishing Group, Apr. 19, 1974; vol. 248, pp. 698-701.

Kim M et al., "Lectin-induced apoptosis of tumour cells" Glycobiology. Oct. 1993 vol. 3 No. 5: pp. 447-453.

Liu K et al., "Chemical Modulation of Cell Fate in Stem Cell Therapeutics and Regenerative Medicine" Cell Chemical Biology; Aug. 18, 2016; 23(8):pp. 893-916.

Palacios R; "ConA triggers T lymphocytes by directly interacting with their receptors" Journal of Immunology. Jan. 1982; vol. 128(1); pp. 337-342.

Pusztai et al., Plant Lectins, 1991, Cambridge University Press, Cambridge, UK, summary only.

Sanford GL et al.; "Stimulation of vascular cell proliferation by beta-galactoside specific lectins." FASEB J. Aug. 1990; 4(11):2912-8)

Sharma et al., "Quantification of Epithelial Cell Differentiation in Mammary Glands and Carcinomas from DMBA- and MNU-Exposed Rats;" PLoS One. 2011; 6(10):e26145.

Smits BM et al., "The Gene Desert Mammary Carcinoma Susceptibility Locus Mcs1a Regulates Nr2f1 Modifying Mammary Epithelial Cell Differentiation and Proliferation" PLoS Genet. Jun. 2013; vol. 9; Issue 6 :e1003549.

Vehmeyer K et al., "Lectin induced increase in clonogenic growth of haematopoietic progenitor cells" European Journal of Haematology. Jan. 1, 1998; vol. 60(1):pp. 16-20.

(Continued)

Primary Examiner — Bridget E Bunner (74) Attorney, Agent, or Firm — D'Ambrosio & Menon, PLLC; Usha Menon

(57) ABSTRACT

A method of treatment of disease condition resulting from deficiency of hematopoietic stem and progenitor cells and/or epidermal stem cells in skin comprising: administering to the subject suffering from said disease conditions, an effective amount of Jack bean lectin or a pharmaceutically acceptable salt thereof for increasing the abundance of hematopoietic stem cells and progenitor cells in bone marrow and/or epidermal stem cells in skin in vivo. Method of treating said conditions in a subject involves administering a single injection of lectin in a dose range of 0.5-2.0 mg/kg body weight to mouse leading to 2.815 to 3.55 fold increase in the abundance of epidermal stem cells in skin and a dose range of 8.0-10.0 mg/Kg body weight to a mouse leading to 2.36 to 6.67-fold increase in the abundance of hematopoietic stem and progenitor cells in bone marrow.

6 Claims, 3 Drawing Sheets

Specification includes a Sequence Listing.