Published online 2017 July 23.

Letter

Prospect of Mesenchymal Stem Cells Conditioned Medium in Tissue Regeneration Tayebeh Sanchooli^{1,*}

¹Department of Anatomical Sciences, School of Medicine, Zahedan University of Medical Sciences, Zahedan, IR Iran

Corresponding author: Tayebeh Sanchooli, Department of Anatomical Sciences, Zahedan, Iran. Tel/Fax: +98-5433295744, E-mail: t_sanchooli@yahoo.com

Received 2017 May 01; Revised 2017 June 04; Accepted 2017 July 02.

Keywords: Mesenchymal Stem Cell, Conditioned Media, Growth Factor, Tissue Regeneration

Dear Editor,

Mesenchymal stem cells (MSCs) are an appealing candidate for tissue regeneration due to their ability to differentiate into many cell types (1). However, the use of stem cells has several problems such as complicated safety and quality management and high cost and difficult cell handling (2). On the other hand, MSCs secrete various trophic factors such as cytokines and growth factors that may regenerate damaged tissues (3). Due to limited survival and differentiation of transplanted MSCs, it is proposed that paracrine effect of MSC secretion is the principal mechanism for tissue regeneration (4).

Some studies investigated the effect of MSC conditioned medium (MSC-CM) on various diseases/conditions such as wound healing and cardiac and ischemic diseases (5). The therapeutic effects of MSC-CM are due to the panel of growth factors and cytokines presented in it. These factors inhibit apoptosis and improve cell proliferation (6). Also, MSC-CM stimulates mobilization and homing of endogenous stem cells into the local site of injury, without the necessity for cell transplantation (2).

Administration of MSC-CM improved bone healing in the rat model (7). It is shown that MSC-CM has a high potential for bone healing, mediated by cytokines such as TGF- β 1, IGF-1, VEGF, and BMP-1. These factors regulate osteogenesis process, angiogenesis, cell migration, and osteoblast proliferation and differentiation (8).

Another application of MSC-CM is the neurological disorders. Some researchers reported that MSC-CM enhances motor activity after spinal cord injury. They concluded that MSC-CM reduced neuronal apoptosis and stimulated angiogenesis. These effects are attributed to some factors in MSC-CM such as NGF and BDNF (9).

Also, the therapeutic effect of MSC-CM was evaluated in the arthritis model. The obtained results showed that MSC-

CM application decreased cartilage destruction and suppressed immune responses by creating a balance between pro- and anti-inflammatory cytokines (10). The application of MSC-CM has several advantages compared with the stem cells such as easily CM package and transportation (5).

In conclusion, the application of MSC-CM in regenerative medicine opened a new window to treat many clinical disorders. This procedure can establish a novel technique in tissue regeneration without the need for stem cells application.

References

- Sun J, Zhou H, Deng Y, Zhang Y, Gu P, Ge S, et al. Conditioned medium from bone marrow mesenchymal stem cells transiently retards osteoblast differentiation by downregulating runx2. *Cells Tissues Organs*. 2012;**196**(6):510–22. doi: 10.1159/000339245. [PubMed: 22906827].
- Osugi M, Katagiri W, Yoshimi R, Inukai T, Hibi H, Ueda M. Conditioned media from mesenchymal stem cells enhanced bone regeneration in rat calvarial bone defects. *Tissue Eng Part A*. 2012;**18**(13-14):1479–89. doi: 10.1089/ten.TEA.2011.0325. [PubMed: 22443121].
- Kim HO, Choi SM, Kim HS. Mesenchymal stem cell derived secretome and microvesicles as a cell free therapeutics for neurodegenerative disorders. *Tissue Eng Regen Med.* 2013;10(3):93–101. doi: 10.1007/s13770-013-0010-7.
- Horie M, Choi H, Lee RH, Reger RL, Ylostalo J, Muneta T, et al. Intraarticular injection of human mesenchymal stem cells (MSCs) promote rat meniscal regeneration by being activated to express Indian hedgehog that enhances expression of type II collagen. Osteoarthritis Cartilage. 2012;20(10):1197–207. doi: 10.1016/j.joca.2012.06.002. [PubMed: 22750747].
- Pawitan JA. Prospect of stem cell conditioned medium in regenerative medicine. *BioMed Res Int*. 2014;2014:1–14. doi: 10.1155/2014/965849.
- Wang CY, Yang HB, Hsu HS, Chen LL, Tsai CC, Tsai KS, et al. Mesenchymal stem cell-conditioned medium facilitates angiogenesis and fracture healing in diabetic rats. *J Tissue Eng Regen Med.* 2012;6(7):559–69. doi: 10.1002/term.461. [PubMed: 21916015].
- Sanchooli T, Norouzian M, Ardeshirylajimi A, Ghoreishi SK, Abdollahifar MA, Nazarian H, et al. Adipose derived stem cells conditioned media in combination with bioceramic collagen scaffolds improved

Copyright © 2017, Gene, Cell and Tissue. This is an open-access article distributed under the terms of the Creative Commons Attribution-NonCommercial 4.0 International License (http://creativecommons.org/licenses/by-nc/4.0/) which permits copy and redistribute the material just in noncommercial usages, provided the original work is properly cited.

calvarial bone healing in hypothyroid rats. *Iran Red Crescent Med J.* 2017;**19**(5). doi: 10.5812/ircmj.45516.

- Linero I, Chaparro O. Paracrine effect of mesenchymal stem cells derived from human adipose tissue in bone regeneration. *PLoS One*. 2014;9(9):107001. doi: 10.1371/journal.pone.0107001. [PubMed: 25198551].
- 9. Cantinieaux D, Quertainmont R, Blacher S, Rossi L, Wanet T, Noel A, et al. Conditioned medium from bone marrow derived mesenchymal

stem cells improves recovery after spinal cord injury in rats: an original strategy to avoid cell transplantation. *PLoS One*. 2013;**8**(8):69515. doi: 10.1371/journal.pone.0069515. [PubMed: 24013448].

 Kay AG, Long G, Tyler G, Stefan A, Broadfoot SJ, Piccinini AM, et al. Mesenchymal Stem Cell-Conditioned Medium Reduces Disease Severity and Immune Responses in Inflammatory Arthritis. *Sci Rep.* 2017;7(1):18019. doi: 10.1038/s41598-017-18144-w. [PubMed: 29269885].