

- Stein J. Y., Levin Y., Uziel O., Abumock H., Solomon Z.* 2018. Traumatic stress and cellular senescence: The role of war-captivity and homecoming stressors in later life telomere length. *J. Affect Disord.* V. 238. P.129.
- Tarte K., Gaillard J., Lataillade J.J., Fouillard L., Becker M., Mossafa H., Tchirkov A., Rouard H., Henry C., Spingard M., Dulong J., Monnier D., Gourmelon P., Gorin N.C., Sensebé L. on behalf of Société Française de Greffe de Moelle et Thérapie Cellulaire.* 2010. Clinical-grade production of human mesenchymal stromal cells: occurrence of aneuploidy without transformation. *Blood.* V. 115. P. 1549.
- Terunuma A., Ashiba K., Takane T., Sakaguchi Y., Terunuma H.* 2019. Comparative transcriptomic analysis of human mesenchymal stem cells derived from dental pulp and adipose tissues. *J. Stem Cells Regen. Med.* V. 15. P. 8.
- Turinetto V., Vitale E., Giachino C.* 2016. Senescence in Human Mesenchymal Stem Cells: Functional Changes and Implications in Stem Cell-Based Therapy. *Int. J. Mol. Sci.* 17. pii: E1164. <https://doi.org/10.3390/ijms17071164>
- Truong N.C., Bui K.H., Van Pham P.* 2018. Characterization of senescence of human adipose-derived stem cells after long-term expansion. *Adv. Exp. Med. Biol.* V. 1084. P. 109. https://doi.org/10.1007/5584_2018_235
- Vassilieva I.O., Reshetnikova G.F., Shatrova A.N., Tsupkina N.V., Kharchenko M.V., Alekseenko L.L., Nikolsky N.N., Burova E.B.* 2018. Senescence-messaging secretome factors, trigger premature senescence in human endometrium-derived stem cells. *Biochem. Biophys. Res. Commun.* V. 496. P. 1162.
- Vigetti D., Moretto P., Viola M., Genassetti A., Rizzi M., Karousou E., Pallotti F., De Luca G., Passi A.* 2006. Matrix metalloproteinase 2 and tissue inhibitors of metalloproteinases regulate human aortic smooth muscle cell migration during in vitro aging. *FASEB J.* V. 20. P. 1118.
- Vulcano F., Milazzo L., Ciccarelli C., Eramo A., Sette G., Mauro A., Macioce G., Martinelli A., La Torre R., Casalbore P., Hassan H.J., Giampaolo A.* 2016. Wharton's jelly mesenchymal stromal cells have contrasting effects on proliferation and phenotype of cancer stem cells from different subtypes of lung cancer. *Exp. Cell Res.* V. 345. P. 190.
- Wagner W., Horn P., Castoldi M., Diehlmann A., Bork S., Saffrich R., Benes V., Blake J., Pfister S., Eckstein V., Ho A.D.* 2008. Replicative senescence of mesenchymal stem cells: a continuous and organized process. *PLoS One.* 3: e2213. <https://doi.org/10.1371/journal.pone.0002213>
- Wang Y., Huso D.L., Harrington J., Kellner J., Jeong D.K., Turney J., McNiece I.K.* 2005. Outgrowth of a transformed cell population derived from normal human BM mesenchymal stem cell culture. *Cyotherapy.* V. 7. P. 509.
- Wang D., Jang D.J.* 2009. Protein kinase CK2 regulates cytoskeletal reorganization during ionizing radiation-induced senescence of human mesenchymal stem cells. *Cancer Res.* V. 69. P. 8200.
- Yang C., Chen Y., Zhong L., You M., Ya Z., Luo M., Zhang B., Yang B., Chen Q.* 2019. Homogeneity and heterogeneity of biological characteristics in mesenchymal stem cells from human umbilical cords and exfoliated deciduous teeth. <https://doi.org/10.1139/bcb-2019-0253>
- Yin H., Pickering J. G.* 2016. Cellular senescence and vascular disease: novel routes to better understanding and therapy. *Can. J. Cardiol.* V. 32. P. 612.
- Yu J., Shi J., Zhang Y., Zhang Y., Huang Y., Chen Z., Yang J.* 2018. The replicative senescent mesenchymal stem / stromal cells defect in DNA damage response and anti-oxidative capacity. *Int. J. Med. Sci.* V. 15. P. 771.
- Zachar L., Bačenková D., Rosocha J.* 2016. Activation, homing, and role of the mesenchymal stem cells in the inflammatory environment. *J. Inflamm. Res.* V. 9. P. 231.
- Zaman W. S., Makpol S., Sathapan S., Chua K. H.* 2014. Long-term *in vitro* expansion of human adipose-derived stem cells showed low risk of tumorigenicity. *J. Tissue Eng. Regen. Med.* V. 8. P. 67.
- Zhang X., Soda Y., Takahashi K., Bai Y., Mitsuru A., Igura K., Satoh H., Yamaguchi S., Tani K., Tojo A., Takahashi T.A.* 2006. Successful immortalization of mesenchymal progenitor cells derived from human placenta and the differentiation abilities of immortalized cells. *Biochem. Biophys. Res. Commun.* V. 351. P. 853.
- Zhang M., Wang Z., Zhao Y., Zhang L., Xu L., Cao L., He W.* 2018. The effect of age on the regenerative potential of human eyelid adipose-derived stem cells. *Stem Cells Int.* 2018. 5654917. <https://doi.org/10.1155/2018/5654917>

CELLULAR AND MOLECULAR CHARACTERISTICS OF REPLICATIVE SENESCENCE OF HUMAN MESENCHYMAL STEM CELLS

D. E. Bobkov^{a, b, *} and G. G. Poljanskaya^a

^a*Institute of Cytology RAS, St. Petersburg, 194064 Russia*

^b*Smorodintsev Research Institute of Influenza, St. Petersburg, 197376 Russia*

*e-mail: bobkov@incras.ru

The widespread use of human mesenchymal stem cells (MSCs) in biomedical technologies necessitates a detailed study of the various properties characteristic of these cells at different periods of their life. The review is devoted to the general and comparative characteristics of different human MSCs, the analysis of various cellular processes accompanying the replicative senescence of MSCs, which is an integral part of the vital activity of these cells during long-term cultivation. Cytogenetic instability, as well as some molecular mechanisms involved in the process of replicative aging of cells, including the reorganization of the extracellular matrix and structures of the cytoskeleton, are considered in more detail in the review.

Keywords: mesenchymal stem cells, replicative senescence, karyotype, extracellular matrix, cytoskeleton