



Letter to the Editor:

Human adult stem cells from menstrual blood and endometrial tissue

Julie ALLICKSON¹, Charlie XIANG^{†‡2,3}

⁽¹⁾*Cryo-Cell International Inc., Oldsmar, FL 34677, USA*

⁽²⁾*State Key Laboratory for Diagnosis and Treatment of Infectious Diseases, the First Affiliated Hospital, School of Medicine, Zhejiang University, Hangzhou 310003, China*

⁽³⁾*S-Evans Biosciences, Hangzhou 311121, China*

[†]E-mail: cxiang@zju.edu.cn

Received Mar. 1, 2012; Revision accepted Mar. 19, 2012

Crosschecked Mar. 29, 2012

doi:10.1631/jzus.B1200062

Document code: A

CLC number: R329.2

The article “Plasticity of human menstrual blood stem cells derived from the endometrium” by Lin *et al.* (2011), published in *Journal of Zhejiang University-SCIENCE B (Biomedicine & Biotechnology)*, described a newly identified mesenchymal-like stem cell (MSC) from human menstrual blood known as MenSC. Here we describe the latest findings in this area and clarify the difference between human adult stem cells from menstrual blood and endometrial tissue.

MenSCs express pluripotent markers such as octamer-binding transcription factor 4 (Oct-4), stage-specific embryonic antigen 4 (SSEA-4), Nanog, and c-kit, in addition to typical MSC surface markers including cluster of differentiation 9 (CD9), CD29, CD44, CD49f, CD90, CD105, and CD166 (Patel *et al.*, 2008; Borlongan *et al.*, 2010). MenSCs have the ability to differentiate into mesoderm tissues such as cartilage (40%–50%), adipose (60%–70%), and bone (45%), with the degree of differentiation similar to or better than bone marrow-derived MSCs. MenSCs also have the capacity to differentiate into neural and cardiac lineages (Patel *et al.*, 2008). Our animal

studies show that the human menstrual blood stem cells can repair several types of damaged cells in vivo (unpublished data).

Meng *et al.* (2007) and Murphy *et al.* (2008) described the cells from menstrual blood termed endometrial regenerative cells (ERCs) that express CD9, CD29, CD44, CD59, CD73, CD90, CD105, and Oct-4, but lack SSEA-4, Nanog, and c-kit. These cells have been demonstrated to differentiate into nine different cell lineages (cardiomyocytic, respiratory epithelial, neurocytic, myocytic, endothelial, pancreatic, hepatic, adipocytic, and osteogenic).

Another source of human adult stem cell from endometrial tissue called eMSC expresses similar MSC markers (Gargett and Masuda, 2010). Both MenSC and eMSC are negative for hematopoietic and endothelial markers such as CD34, CD38, CD45, and CD133 (Patel *et al.*, 2008; Gargett and Masuda, 2010). Major differences between the cells include the invasive process required to retrieve the eMSC as well as the growth properties identified in vitro where eMSC but not MenSC requires growth factors and an extracellular matrix. Independent studies comparing eMSC to MenSC demonstrated 57 significant and differentially expressed genes (unpublished data).

Both MenSC and eMSC are similar to bone marrow-derived MSC; however, stromal-derived factor-1 (STRO-1), a marker used to prospectively isolate bone marrow-derived MSC (Gronthos *et al.*, 2003), has not been found in cells from the endometrial tissue (Gargett and Masuda, 2010) or in MenSCs, although decidual cells are known to express STRO-1 (García-Pacheco *et al.*, 2001).

MenSCs recapitulate pluripotency properties as an alternative adult stem cell source that circumvents ethical and logistical limitations of embryonic stem (ES) cells. Recent studies reveal great potential of

[†] Corresponding author

MenSCs for treating stroke (Borlongan *et al.*, 2010) and liver damage (unpublished data).

References

- Borlongan, C.V., Kaneko, Y., Maki, M., Yu, S.J., Ali, M., Allickson, J.G., Sanberg, C.D., Kuzmin-Nichols, N., Sanberg, P.R., 2010. Menstrual blood cells display stem cell-like phenotypic markers and exert neuroprotection following transplantation in experimental stroke. *Stem Cells Dev.*, **19**(4):439-452. [doi:10.1089/scd.2009.0340]
- García-Pacheco, J.M., Oliver, C., Kimatrai, M., Blanco, F.J., Olivares, E.G., 2001. Human decidual stromal cells express CD34 and STRO-1 and are related to bone marrow stromal precursors. *Mol. Hum. Reprod.*, **7**(12):1151-1157. [doi:10.1093/molehr/7.12.1151]
- Gargett, C.E., Masuda, H., 2010. Adult stem cells in the endometrium. *Mol. Hum. Reprod.*, **16**(11):818-834. [doi:10.1093/molehr/gaq061]
- Gronthos, S., Zannettino, A.C., Hay, S.J., Shi, S., Graves, S.E., Kortessidis, A., Simmons, P.J., 2003. Molecular and cellular characterisation of highly purified stromal stem cells derived from human bone marrow. *J. Cell Sci.*, **116**(9):1827-1835. [doi:10.1242/jcs.00369]
- Lin, J., Xiang, D., Zhang, J.L., Allickson, J., Xiang, C., 2011. Plasticity of human menstrual blood stem cells derived from the endometrium. *J. Zhejiang Univ.-Sci. B (Biomed. & Biotechnol.)*, **12**(5):372-380. [doi:10.1631/jzus.B1100015]
- Meng, X., Ichim, T.E., Zhong, J., Rogers, A., Yin, Z., Jackson, J., Wang, H., Ge, W., Bogin, V., Chan, K.W., *et al.*, 2007. Endometrial regenerative cells: a novel stem cell population. *J. Transl. Med.*, **5**(1):57. [doi:10.1186/1479-5876-5-57]
- Murphy, M.P., Wang, H., Patel, A.N., Kambhampati, S., Angle, N., Chan, K., Marleau, A.M., Pyszniak, A., Carrier, E., Ichim, T.E., *et al.*, 2008. Allogeneic endometrial regenerative cells: an "Off the shelf solution" for critical limb ischemia? *J. Transl. Med.*, **6**:45. [doi:10.1186/1479-5876-6-45]
- Patel, A.N., Park, E., Kuzman, M., Benetti, F., Silva, F.J., Allickson, J.G., 2008. Multipotent menstrual blood stromal stem cells: isolation, characterization, and differentiation. *Cell Transplant.*, **17**(3):303-311. [doi:10.3727/096368908784153922]

2010 JCR of Thomson Reuters for JZUS-A and JZUS-B

ISI Web of Knowledge SM									
Journal Citation Reports [®]									
WELCOME		HELP		RETURN TO LIST		2010 JCR Science Edition			
Journal: Journal of Zhejiang University-SCIENCE A									
Mark	Journal Title	ISSN	Total Cites	Impact Factor	5-Year Impact Factor	Immediacy Index	Citable Items	Cited Half-life	Citing Half-life
<input type="checkbox"/>	J ZHEJIANG UNIV-SC A	1673-565X	442	0.322		0.050	120	3.7	7.1
Journal: Journal of Zhejiang University-SCIENCE B									
Mark	Journal Title	ISSN	Total Cites	Impact Factor	5-Year Impact Factor	Immediacy Index	Citable Items	Cited Half-life	Citing Half-life
<input type="checkbox"/>	J ZHEJIANG UNIV-SC B	1673-1581	770	1.027		0.137	124	3.5	7.5