Colloquia in Cellular Signaling

Venue: Medical University Vienna, Center for Physiology and Pharmacology, Institute of Pharmacology, Waehringerstrasse 13a, 1090 Vienna, "Leseraum". (Sonja Sucic, Tel.: (01) 40160-31371, E-mail: sonja.sucic@meduniwien.ac.at)

Friday 06.10.2017 11:00 h Host: Eva Zebedin

Towards an artificial stem cell niche: biomaterials for directing hematopoietic stem cell behavior

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Abstract:

Blood is replenished with billions of fresh cells every day throughout the entire life span. The source of these cells are the so-called hematopoietic (= blood forming) stem cells (HSCs). Their ability to reconstitute the entire blood system makes them the key to the cure of many hematological diseases. Upon transplantation from a healthy donor, they are able to replace the diseased hematopoietic system of the patient. However, this treatment is restricted by the limited availability of HSCs. To overcome that limitation, controlling HSC behavior in terms of proliferation or differentiation in vitro, is an important goal of nowadays HSC research. *In vivo* HSCs are controlled by a highly specialized microenvironment – the niche – within the bone marrow. In this niche HSCs are supported by mutual cell-cell as well as cellmatrix interactions. While it is clear that biological and/or chemical parameters play an important role in this interplay, surprisingly little attention was paid to physical signals that are transmitted by the niche microenvironment. In the last years, we found that these physical signals include matrix stiffness, nanostructure as well as the three-dimensional architecture. In reductionist approaches, in which we studied only one parameter at a time, we could show that all of these parameters impact HSC behavior. In order to achieve the goal of a synthetic stem cell niche to guide HSC behavior, the complexity of the natural HSC niche, which combines a variety of different signals, has to be taken into account. For this purpose, we increased the complexity of our systems to study the synergistic effects of different biological and/or physical signals. With these experiments we hope to get one step closer towards a synthetic stem cell niche that is as simple as possible but as complex as necessary to instruct HSCs.