

8th International Conference on
TISSUE SCIENCE AND REGENERATIVE MEDICINE
September 11- 12, 2017 Singapore

Cellular aging of the immune system: Causes and consequences

Renu Balyan^{1,4}, Rupali Gund¹, Amanpreet Singh Chawla¹, Saurabh Pradhan², Satyajeet Khare², Sanjeev Galande², Jeannine Durdik³, Anna George¹, Vineeta Bal¹ and Satyajit Rath¹

¹National Institute of Immunology, India

²Indian Institute of Science Education and Research, India

³University of Arkansas, USA

⁴National University of Singapore, Singapore

Functional heterogeneity in apparently homogenous T-cell populations has been well reported. However, what factors lead to such variations within a T-cell population is still unclear. Depending upon the time of egress from the thymus and duration of residence in the peripheral lymphoid organs, individual cells would be of relatively different ages, however, consequences of such cellular aging have not been addressed as there are no phenotypic markers to estimate the age of individual naive T-cells. We show that CD8 levels reduce over time, as a result of MHC-I mediated tonic signals and lower CD8 levels can be used as a surrogate marker to identify relatively aged cells in a naive CD8 T-cell population. We observed that naive CD8 T-cells from aged mice have lower CD8 levels than naive CD8 T-cells of young mice. Further, when naive T-cells from young mice are separated into CD8-hi and CD8-lo subsets, we find that CD8-lo cells are smaller, respond poorly and are more susceptible to death. Bulk mRNA sequencing showed that peripheral MHC-I-mediated tonic signaling leads to post-thymic age-associated differences in CD8 T-cells and major changes in their signaling, transcriptional and functional landscapes. Our findings also suggest the feasibility of potential pharmacological interventions for improved T-cell responses during vaccination of older people via either anti-oxidant or DUSP (Dual-specificity phosphatase) inhibitor small molecules.

zhuihengzhu@163.com