

Early thymic selection and conditioning begin the shaping of the T cell repertoire.

Susannah Barbee, Martin Woodward, Richard Lifton, Robert Tigelaar, **Adrian Hayday**
Guy's Hospital, London, England, United Kingdom

Thymocyte differentiation is the route by which a diverse, functionally pleiotropic peripheral T cell compartment can be established. We have sought to identify molecular interactions that shape thymocyte development, and will discuss novel immunoglobulin superfamily genes, known as *Skint* genes, that are expressed by thymic epithelium and that select the repertoire of “unconventional T cells” that contribute to “transitional immunity”. By contrast to “stress-activated” immune sentinels, these genes appear to denote normality. We shall also consider molecular interactions of early thymocytes with other thymocytes; how this promotes heterogeneity among thymocytes; and how this may predispose cells to certain functional competences. The integration of these events begins the shaping of the T cell repertoire.

References:

Pennington, D.J., Vermijlen, D., Wise, E.L., Clarke, S.L., Tigelaar, R.E., and Hayday, A.C. “The Integrated Responses of Conventional and Unconventional T cells” *Adv in Immunology* 87:27-59

Lewis, J.M., M.Girardi, S.J. Roberts, S. Barbee, A. C. Hayday & R.E. Tigelaar “Selection of the cutaneous intraepithelial gamma delta T cell repertoire by a thymic stromal determinant” *Nature Immunol.* 8:843-50

Silva-Santos B, Pennington DJ, Hayday AC “Lymphotoxin-Mediated Regulation of $\gamma\delta$ Cell Differentiation by $\alpha\beta$ T Cell Progenitors” *Science.* 307:925-928

Hayday AC and Pennington, DP. “Key factors in the organized chaos of early T cell development” *Nature Immunol.* 8:137-44

Pennington DJ, Silva-Santos B, Silberzahn T, Escorcio-Correia M, Woodward MJ, Roberts SJ, Smith AL, Dyson PJ, Hayday AC. “Early events in the thymus affect the balance of effector and regulatory T cells.” *Nature* 444: 1073-1077