The Interplay between Cellular and Humoral Innate Immunity

MANTOVANI, Alberto

Scientific Director, Istituto Clinico Humanitas and President, Humanitas Foundation for

Research, Italy

Email: alberto.mantovani@humanitas.it

ABSTRACT

Innate immunity consists of a cellular and a humoral arm. The role of negative regulators, including TIR8 (a member of the IL-1 receptor family) and D6 (a decoy and scavenger chemokine receptor) will be discussed. We have used the long pentraxin PTX3 as a paradigm for the humoral arm of innate immunity and its interplay with cells. PTX3 is a multifunctional soluble pattern recognition receptor characterized by a C-terminal domain highly homologous to C-reactive protein and serum amyloid P component, associated to a N-terminal domain unrelated to other known proteins. PTX3 is produced upon stimulation with proinflammatory cytokines and Toll-like receptor engagement by different cell types, including endothelial cells, monocytes/macrophages, dendritic cells, fibroblasts and epithelial cells.

The molecule binds with high affinity complement component, microbial moieties, growth factors and apoptotic cells. PTX3 plays non-redundant functions including innate immunity against selected microorganisms to regulation of inflammation. In addition PTX3 has a regulatory function. PTX3 serves as a paradigm for the interplay between the cellular and the humoral arm of innate immunity.

Bottazzi B, Doni A, Garlanda C, Mantovani A. An Integrated View of Humoral Innate Immunity: Pentraxins as a Paradigm. Annu Rev Immunol. 2010, 28:157-83.

Deban L, Russo RC, Sironi M, Moalli F, Scanziani M, Zambelli V, Cuccovillo I, Bastone A, Gobbi M, Valentino S, Doni A, Garlanda C, Danese S, Salvatori G, Sassano M, Evangelista V, Rossi B, Zenaro E, Constantin G, Laudanna C, Bottazzi B, Mantovani A. Regulation of leukocyte recruitment by the long pentraxin PTX3. Nat Immunol. 2010, 11:328-34.

Sica A, Mantovani A.

Macrophage plasticity and polarization: in vivo veritas. J Clin Invest. 2012, 122:787-95.