

Immunology essay questions

Discuss the roles of innate immunity in antigen recognition by T cells during responses to infection.

T cells are categorised into two main classes depending on their expression of CD 4 and CD8 cell surface proteins. Dependant on the type of T cell, during antigen recognition CD 4 or CD8 cells associate with the T cell receptor (TCR) and bind to conserved regions of major histocompatibility complex (MHC) class I/II molecules. The co-stimulation of the singular MHC molecule by TCR and its CD 4/CD8 coreceptor, augments the avidity of TCR binding to its target cell.

Dendritic cells (DCs) directly link the innate immune system with the adaptive immune system. There are two main functional types of DCs; conventional DCs (cDCs) and plasmacytoid DCs (pDCs). Both types are derived from hematopoietic bone marrow progenitor cells. DCs are the only kind of cell type capable of presenting antigen to activate naive T cells in the innate immune response. DCs take up antigen through innate immune receptors (PRRs) and internally process them to create short peptide segments that are loaded onto MHC class I and II. DCs then migrate to the secondary lymphoid organs so they can present antigens to T cells, therefore instigating antigen-specific immune responses, or immunological tolerance.

The capability of DCs to regulate innate immunity is reliant on DC maturation. After DCs uptake and process the antigen, several factors can induce their maturation, including whole bacteria/bacterial-derived antigens (e.g., lipopolysaccharide, LPS), inflammatory cytokines, ligation of selective cell surface receptors (e.g., CD40) and viral products (e.g., double-stranded RNA). During DC maturation cells become less phagocytic and endocytic, there is also increased DC proliferation as well as upregulation of cell surface protein and cytokine expression (to optimize antigen presentation to T cells). During maturation DCs also secrete type 1 interferons which inhibit pathogen replication, induce apoptosis of infected cells, and again enhance antigen presentation. Antigen presentation is also enhanced through the action of type 1 IFNs. NK cells and T cells secrete cytokines like IFN γ which enhance antigen presentation of DCs.

Macrophages are sentinel cells which are also capable of presenting antigen to T cells, but are unable to initiate a primary immune response, unlike DCs which are optimal in antigen presentation.

Studies have shown that in aged C57BL/6 mice that were susceptible to the mousepox virus, DCs were dysfunctional and unable to activate NK cells. In an NK-sensitive tumour model, this defect resulted in failure to eliminate RMA-S lymphoma mutant tumour cells (Guo Z, *et al* 2014). This shows the key role of active DCs in the innate immune system, as aged DCs have a lessened ability to induce antigen-specific T cells, leading to deficient tumour immunity. In addition, LAV models (combined with different viral models) have shown that DCs represent an important family of cells promoting antiviral innate immunity (Summerfield A, *et al* 2009).

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