

1. Introduction to Immunology

- 1.1. A historical perspective of immunology.
- 1.2. The basics of hematopoiesis, focused on the development of myeloid, lymphoid and erythroid cells. The categories of cells types in terms of characteristics and function in an immune response.
- 1.3. The anatomy of the lymphoid organs, tissues, and the movement of leukocytes in the body.
- 1.4. General properties of receptor-ligand interactions.
- 1.5. Overview of signaling molecules and pathways.

2. Innate Immune Responses

- 2.1. Overview of the characteristics and functions of the innate immune system.
- 2.2. Functional elements of innate immunity: barriers, cells, tissues, soluble mediators.
- 2.3. Pattern recognition receptors, pathogen (damage)-associated molecular patterns.
- 2.4. Activation of innate immune cells and signaling pathways.
- 2.5. Mechanisms of extravasation, phagocytosis, and pathogen destruction.
- 2.6. Function and components of the inflammatory response and the subsequent outcomes of localized and systemic inflammation.
- 2.7. Acute phase response and the resulting systemic changes in the body.
- 2.8. Complement system components, activation and regulation of the complement system and its role in the recognition and elimination of pathogens, cooperation with the inflammatory response and hemostatic/coagulation pathways.
- 2.9. The role of NK cells and interferons in the recognition and elimination of virally-infected cells.
- 2.10. Characteristics and functions of innate lymphoid cells.
- 2.11. Processes performed by innate immune cells that result in the activation of T and B lymphocytes to a specific pathogen.

3. Lymphocyte Development and Activation

- 3.1. Organization and expression of lymphocyte receptor genes.
- 3.2. Discriminate between BCR and TCR gene rearrangements, functional parts of each receptor, cell signaling events through these receptors, and cell surface marker expression during development and on mature lymphocytes.
- 3.3. B and T cell development from a stem cell to a mature cell.
- 3.4. Positive and negative selection of T cells in the thymus contrasted with positive and negative selection of B cells in the bone marrow, along with the role of other secondary lymphoid organs during these developmental processes.
- 3.5. Genetic organization of the Major Histocompatibility Complex (MHC) including the concepts of co-dominant expression, polygeny, polymorphism, and linkage disequilibrium.
- 3.6. Structure, function, and expression patterns of MHC Class I and MHC Class II proteins.
- 3.7. The role of antigen presenting cells and antigen processing and presentation pathways in T lymphocyte activation.
- 3.8. Mechanisms of B cell activation, through both T-independent and T-dependent antigens.
- 3.9. Antibody structure, structural isoforms, isotypes, antigen-antibody interactions.

4. Humoral and Cell-mediated Immune Responses

- 4.1. T-independent B cell responses
- 4.2. T-dependent B cell responses, linked recognition
- 4.3. The role of T cells, co-receptors and cytokines in the processes of isotype switching, somatic hypermutation, affinity maturation, and the development of memory versus plasma B cells.
- 4.4. Antibody-mediated effector functions.
- 4.5. The role of antigen presenting cells (APCs) and cytokines on the differentiation of T cell subsets, and the development of memory T cells.
- 4.6. CD4+ T cell-mediated functions.
- 4.7. CD8+ T, NK, and NKT functions.
- 4.8. Regulation of humoral and cell-mediated responses.
- 4.9. Characteristics and function of memory responses.

5. Mucosal Immunity

- 5.1. The cells and tissues of the mucosal immune system, with emphasis on the gastrointestinal tract.
- 5.2. Prominent species of the human microbiome and the contribution of non-bacterial species to the microbiome: mycobiome and virome
- 5.3. The ability of the mucosal immune system to switch from a tolerant response to an active immune response against pathogens
- 5.4. Characteristics of the mucosal immune system and the microbiome that facilitate the maintenance of intestinal homeostasis
- 5.5. Microbial dysbiosis and alterations in mucosal immune components that contribute to aberrant immune responses and disease.

6. Immunopathologies

- 6.1. Hypersensitivities
 - 6.1.1. Characteristics of Type I hypersensitivity reactions, roles of immune components contributing to Type I HS and clinical outcomes – allergy, asthma (driven by Th2/Th17), anaphylactic shock, food allergy/tolerance to food antigens, hygiene hypothesis
 - 6.1.2. Similarities of and differences between Type II and Type III hypersensitivity immune mechanisms and clinical outcomes.
 - 6.1.3. Immune components and mechanisms of delayed-type (Type IV) hypersensitivity and clinical outcomes – granuloma formation, contact dermatitis, celiac disease
- 6.2. General principles of immune regulation
 - 6.2.1. Immune dysregulation and chronic inflammation, causes and consequences
- 6.3. Autoimmunity
 - 6.3.1. Mechanisms of central and peripheral tolerance.
 - 6.3.2. Processes that result in a break in tolerance.
 - 6.3.3. Causes of autoimmune diseases, the genetics of autoimmune disease including both MHC and non-MHC links to disease, and the HLA-associate risk factors for these diseases.

UIP Content Objectives

- 6.3.4. The role of humoral versus cell-mediated immune responses in autoimmune disease mechanisms and pathologies.
- 6.3.5. Therapeutic approaches for treating autoimmune disease.
- 6.4. Transplantation
 - 6.4.1. HLA antigens and their role in the organ/tissue transplantation of Auto-, Allo-, Iso-, and Xeno- grafts.
 - 6.4.2. The immunologic mechanisms involved in the rejection of transplanted tissues and organs.
 - 6.4.3. The concept of bone marrow transplantation and the role of the immune response in graft versus host disease (GVHD).
- 6.5. Cancer
 - 6.5.1. Tumor associated antigens and their recognition on cancer cells by the immune system.
 - 6.5.2. Mechanisms used by tumor cells to evade detection and destruction by the immune system.
 - 6.5.3. Cancer-specific immunotherapies, including CAR-T, anti-CTLA4, anti-PD1 and similar checkpoint inhibitors, Tumor-infiltrating lymphocyte (TILs) and other breakthrough technologies used to attack cancer cells.
- 7. Immunodeficiencies**
 - 7.1. Primary immunodeficiencies of the innate and adaptive immune systems and the effect on the elimination of pathogens.
 - 7.2. Pathogens and infectious diseases associated with specific immunodeficiencies.
 - 7.3. Secondary immunodeficiencies of the adaptive immune responses associated with poor nutrition, immunosuppressive drugs, and chronic infections.
 - 7.4. The history of HIV/AIDS, interactions of HIV with immune cells and the consequences of the ongoing immune response, pathogens associated with HIV infection and AIDS.
- 8. Infectious Disease and Vaccines**
 - 8.1. Structural and physiological features used for classification and identification of infectious agents.
 - 8.2. Features of infectious agents that contribute to their pathogenicity.
 - 8.3. Common events that occur in infectious diseases such as host encounter and entry, adherence and colonization, spread, evasion of host defenses, and tissue damage.
 - 8.4. The spectrum of pathogenicity from commensal to obligate pathogens.
 - 8.5. The immune response to viruses.
 - 8.6. The immune response to bacteria.
 - 8.7. The immune response to parasites.
 - 8.8. The immune response to fungi.
 - 8.9. Evasion of innate and adaptive immune mechanisms by pathogens.
 - 8.10. History of vaccination.
 - 8.11. Protective immunity through passive and active immunization.
 - 8.12. Characteristics, strengths and weaknesses of varying vaccine strategies. Memory responses and original antigenic sin.

UIP Content Objectives

- 8.13. Purpose and immunologic outcome of conjugate and multivalent component vaccines and adjuvants.

9. Immunologic Methods and Experimental Systems

- 9.1. Monoclonal and polyclonal antibody generation and applications.
- 9.2. Purpose and use of immunoprecipitation and agglutination reactions.
- 9.3. Enzyme-linked immunosorbent assays
- 9.4. Immunofluorescence-based imaging techniques.
- 9.5. Flow cytometry and cell sorting.
- 9.6. Use of mouse model systems – inbred and congenic strains, adoptive transfer experiments, transgenic mice: knock-in, knockout, cre/lox systems

10. Immunotherapies

- 10.1. History of development
- 10.2. Activating immunotherapies
- 10.3. Suppressing immunotherapies

401 – INNATE IMMUNITY

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- 2.12. Processes performed by innate immune cells that result in the activation of T and B lymphocytes to a specific pathogen.

5. Mucosal Immunity

- 5.1. The cells and tissues of the mucosal immune system, with emphasis on the gastrointestinal tract.
- 5.3. The ability of the mucosal immune system to switch from a tolerant response to an active immune response against pathogens
- 5.4. Characteristics of the mucosal immune system and the microbiome that facilitate the maintenance of intestinal homeostasis
- 5.5. Microbial dysbiosis and alterations in mucosal immune components that contribute to aberrant immune responses and disease.

6. Immunopathologies

- 6.2. General principles of immune regulation
 - 6.2.1. Immune dysregulation and chronic inflammation, causes and consequences

7. Immunodeficiencies

- 7.1. Primary immunodeficiencies of the innate and adaptive immune systems and the effect on the elimination of pathogens.

402 – ADAPTIVE IMMUNITY

Brief review of 2.1, 2.3, 2.11

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403- PATHOGEN-IMMUNE SYSTEM INTERACTIONS

Review 2.5 – 2.9

Review 3.8, 3.9

Review 4.4, 4.6, 4.7, 4.9

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404 – IMMUNE-MEDIATED DISEASES

Review 4.8

Review PLO 9

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UIP Content Objectives Mapped to Courses

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