

Chapter 6

Diseases of Immunity:

1. The innate immune system includes the following components EXCEPT
 - (a) lung surfactant
 - (b) complement
 - (c) Natural killer (NK) cells
 - (d) dendritic cells
 - (e) B lymphocytes

2. Regarding the immune system
 - (a) antibodies play a crucial role in the innate immune system
 - (b) The classical pathway of complement activation is part of adaptive immunity
 - (c) mannose binding lectins are released by microbes, and are important for complement activation
 - (d) C reactive protein is a by-product of various adaptive immunity responses
 - (e) none of the above is true

3. Regarding the immune system
 - (a) T lymphocytes make up 30% of circulating lymphocytes
 - (b) humoral immunity is part of the innate immune response
 - (c) cell mediated immunity provides defence against intracellular organisms
 - (d) NK cells play a pivotal role in cell mediated immunity
 - (e) diverse receptors on lymphocytes show that they are inherently specific for a particular microbe

4. T cells

- (a) are activated by soluble antigen
- (b) have a receptor for antigen (TCR) that is made up of α , β , and γ subunits
- (c) CD8 molecules bind to the class II MHC molecules
- (d) with the $\gamma\delta$ TCR recognise peptides and lipids without the need for antigen presentation
- (e) receptor recognises a very limited number of peptides

5. The position of immune cells in the spleen is (*old paper 2004*)

- (a) T cells in the medullary cords
- (b) B cells in the paracortical regions of the white pulp
- (c) Macrophages in the paracortical regions of the white pulp
- (d) B cells in the perifollicular regions of the white pulp
- (e) Neutrophils cells in the medullary cords

6. What immune cell type predominates in the perifollicular regions of the white pulp in the spleen? (*variation 2004*)

- (a) T lymphocytes
- (b) B Lymphocytes
- (c) Macrophages
- (d) Plasma cells
- (e) Mast cells

7. What receptor is present on the surface of all naïve B cells

- (a) IgM and IgD
- (b) IgM and IgG
- (c) IgM and IgA
- (d) IgG and IgD
- (e) IgG and IgE

8. Regarding the proportion of circulating lymphocytes

- (a) the B cell make up 50%
- (b) the T cell make up 60-70%
- (c) neutrophils make up 90%
- (d) Plasma cells make up 50%
- (e) none of the above figures is correct

9. B lymphocytes

- (a) are the most common circulating lymphocyte
- (b) are activated independently of T helper cells
- (c) are integral to the cell mediated system of immunity
- (d) have a receptor for CMV virus
- (e) may be activated by protein and non-protein antigens

10. Natural killer cells

- (a) are leukocytes
- (b) are a predominant cell in adaptive immunity
- (c) have the ability to kill a cell without prior sensitisation
- (d) unlike other cells of the immune system, do not produce cytokines
- (e) kill cells that over-express class I MHC molecules, such as tumour cells

11. Regarding cytokines

- (a) They are specific for one cell type
- (b) They are often redundant
- (c) Their main role is in innate immunity
- (d) They act exclusively as stimulators for immune responses
- (e) none of the above

The class I major histocompatibility complex (MHC)

- (a) is expressed on all cells of the body
- (b) is encoded for on the short arm of chromosome 16
- (c) has a low level of polymorphism
- (d) are encoded for by three loci designated HLA-A, HLA-B and HLA-C
- (e) β -microglobulin on the MHC is expressed on the HLA A locus

Mast cells

- (a) are bone marrow derived cells
- (b) are the primary cell involved in type II hypersensitivity reactions
- (c) inhibit platelet activation, but are otherwise inflammatory cells
- (d) can phagocytose antigen
- (e) none of the above is true

Regarding mast cells, which statement is false

- (a) They can degranulate with exposure to morphine
- (b) They are activated with cross linking of high affinity IgE Fc receptors
- (c) They can degranulate with exposure to C5a
- (d) Basophils are similar to mast cells, but are in much higher concentrations in certain tissues (eg the lung)
- (e) They can be activated by IL-8

Examples of primary mediators released by Mast cells would be

- (a) cytokines
- (b) leukotrienes
- (c) Platelet activating factor
- (d) heparin
- (e) Prostaglandin D₂

An example of secondary mediators released by Mast cells would be

- (a) histamine
- (b) platelet activating factor
- (c) heparin
- (d) chymase
- (e) tryptase

In systemic anaphylaxis

- (a) the severity of the response is proportional to the concentration of the antigen
- (b) there is widespread oedema, but the larynx is spared
- (c) a previous history of some form of allergy is always present
- (d) the symptoms usually follow administration of foreign proteins
- (e) hives are not a feature

Complement membrane attack complex can be activated by

- (a) IgE
- (b) IgA
- (c) IgD
- (d) IgG
- (e) none of the above

Regarding hypersensitivity reactions

- (a) the presence of antibody-antigen complexes in the circulation implies disease
- (b) Type III hypersensitivity is generally due to reactions against endogenous antigens
- (c) Pemphigus vulgaris is an example of complement-mediated inflammation
- (d) The T_H1 -type helper cell promotes the synthesis of IgE in type I hypersensitivity reactions
- (e) In type III hypersensitivity, immune complexes are typically deposited in vessel walls

In type III hypersensitivity reaction,

- (a) phase 2 begins a week after exposure to antigen
- (b) small or intermediate size antigen-antibody complexes are more likely to cause this disease process
- (c) wherever the complexes deposit, the tissue damage is similar
- (d) phase 3 begins approximately 10 days after exposure to the antigen
- (e) all of the above is true

Central tolerance

- (a) means that in a normal individual, T cells bearing receptors for autoantigens are never present
- (b) prevents immature B cells from reacting to self, as they undergo apoptosis if exposed to membrane bound antigen within the bone marrow
- (c) occurs for B cells in the thymus
- (d) describes a state of anergy towards antigen presenting cells
- (e) none of the above is true

Peripheral tolerance is maintained by all of the following mechanisms except

- (a) Anergy
- (b) deletion of T-cells that express high affinity for self antigens during maturation in the thymus
- (c) clonal deletion by activation-induced cell death
- (d) Suppression by regulatory T cells
- (e) Antigen sequestration

Antigen sequestration is

- (a) the phagocytosis of bacteria by leukocytes
- (b) the removal of self recognising lymphocytes from the lymphocyte population
- (c) where tissues in which antigens are located do not communicate with blood or lymph
- (d) the uptake of antigen by Peyer's patches
- (e) the presentation of antigenic products on MHC II molecules

Concerning the mechanisms of autoimmune disease

- (a) the actual genes associated with most autoimmune diseases is now known
- (b) normal MHC molecules are incapable of presenting self antigens
- (c) few autoimmune diseases are associated with preceding infections
- (d) molecular mimicry may play a role in the development
- (e) these diseases can spontaneously resolve, a process known as epitope spreading

Transmission of HIV

- (a) through needlestick injury is approximately 10%
- (b) through needlestick injury can be reduced eightfold when a patient is given antiretroviral therapy
- (c) in utero is the most common mode of mother-to-infant transmission
- (d) from blood transfusion has been eliminated
- (e) can be transmitted by mosquito

HIV

- (a) is a double stranded RNA virus
- (b) has two viral glycoproteins, gp120 and gp41, which are vital for infection of cells
- (c) envelope is lipopolysaccharide
- (d) has limited variability expressed in its genome
- (e) invades CD4+ cells, but does not invade macrophages

Concerning HIV

- (a) accumulation of T tropic virus is a sign of the final rapid phase of disease progression
- (b) M-tropic strains use CCR5 chemokine receptor to infect host cells
- (c) T-tropic strains use CXCR4 chemokine receptor to infect host cells
- (d) M-tropic viruses are the dominant virus type found in newly infected individuals
- (e) all of the above is true

Concerning HIV infection

- (a) Infected monocytes transport the virus to the brain
- (b) The type of virus that infects the microglia is M-tropic
- (c) Neurons are infected by HIV
- (d) Macrophages are easily destroyed by HIV virus replicating in the phagolysosome
- (e) A and B are true

Concerning HIV complications

- (a) Invasive candidiasis is common
- (b) Atypical pneumonia caused by *Mycoplasma pneumoniae* is one of the sentinel infections
- (c) Toxoplasmosis is responsible for 50% of mass lesions in the CNS of HIV patients
- (d) 5% of infected individuals will develop a malignancy
- (e) Kaposi sarcoma is more common in patients who acquired the disease parenterally

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Answers:

1. The innate immune system includes the following components EXCEPT (p194, fig 6-1)
 - (a) lung surfactant
 - (b) complement
 - (c) Natural killer (NK) cells
 - (d) dendritic cells
 - (e) B lymphocytes**

2. Regarding the immune system p195
 - (a) antibodies play a crucial role in the innate immune system *of the neonate (breast feeding)*
 - (b) The classical pathway of complement activation is part of adaptive immunity**
 - (c) mannose binding lectins *are circulating proteins*, and are important for complement activation when bound to bacterial cell wall
 - (d) *C reactive protein is produced by the liver*, activates C1q, its role is uncertain, and is usually at <1mg/dL in the blood
 - (e) none of the above is true (*wrong*)

3. Regarding the immune system p195-6
 - (a) T lymphocytes make up **60-70%** of circulating lymphocytes
 - (b) humoral immunity is part of the adaptive immune response
 - (c) cell mediated immunity provides defence against intracellular organisms**
 - (d) NK cells play a pivotal role *in innate immunity*
 - (e) the lymphocyte has great diversity of receptors, compared with the innate immune system, but lymphocytes are not inherently specific for a microbe

4. T cells p197-198
 - (a) **B cells are activated by soluble antigen. T cells require presented antigen**
 - (b) have a receptor for antigen (TCR) that is made up of α , β or $\gamma\delta$ subunits
 - (c) CD8 molecules bind to the *class I MHC molecule (cytotoxic)*, *CD4 binds to MHC II (T_H cells)*
 - (d) with the $\gamma\delta$ TCR recognise peptides and lipids without the need for antigen presentation**
 - (e) receptor recognises a *large number of peptides*

5. The position of immune cells in the spleen is (*old paper 2004*)
 - (a) **Plasma cells** in the medullary cords of lymph nodes
 - (b) **T cells** in the paracortical regions of the white pulp
 - (c) **T cells** in the paracortical (arteriolar) regions of the white pulp
 - (d) B cells in the perifollicular regions of the white pulp**
 - (e) **Plasma cells** in the medullary cords of lymph nodes

6. What immune cell type predominates in the perifollicular regions of the white pulp in the spleen? (p523 Ganong 21st, p198-99). (*variation 2004*)
 - (a) T lymphocytes (paracortical)
 - (b) B Lymphocytes**
 - (c) Macrophages
 - (d) Plasma cells
 - (e) Mast cells

7. What receptor is present on the surface of all naïve B cells p198
- (a) **IgM and IgD**
 - (b) IgM and IgG
 - (c) IgM and IgA *IgA is secreted*
 - (d) IgG and IgD *IgG is formed as part of the adaptive immunity and is seen as a marker for immunity*
 - (e) IgG and IgE *IgE is formed in response to multicellular organism infection*
8. Regarding the proportion of circulating lymphocytes p198-99
- (a) the *B cell make up 10-20%*
 - (b) **the T cell make up 60-70%**
 - (c) neutrophils are not a *lymphocyte, but make up 90% of the WCC usually*
 - (d) Plasma cells *are activated B cells, and hence will be a portion of 20%%*
 - (e) none of the above figures is correct (*wrong*)
9. B lymphocytes p199
- (a) *T lymphocytes* are the most common circulating lymphocyte
 - (b) *require help from T helper cells via CD40, which matures the B cell and allows change to IgG production*
 - (c) are integral to the *humoral system* of immunity
 - (d) have a receptor for *EBV virus (CD21), and are easily infected by this virus*
 - (e) **may be activated by protein and non-protein antigens**
10. Natural killer cells p201-202
- (a) are *lymphocytes (10-15% circulating)*
 - (b) are a predominant cell in *innate* immunity
 - (c) **have the ability to kill a cell without prior sensitisation: virally infected, tumour etc**
 - (d) like other cells of the immune system, *produce cytokines which activate $m\phi$, and helps with the differentiation of CD4+ cells*
 - (e) kill cells that *under-express class I MHC molecules*, such as tumour cells
11. Regarding cytokines p202
- (a) They *have multiple* cell type targets
 - (b) **They are often redundant, meaning that their functions overlap**
 - (c) Their role is in *innate, adaptive immunity and inflammatory responses*
 - (d) They act as *stimulators for immune responses, and inhibitory (IL-10, TGF- β)*
 - (e) none of the above (*wrong*)
- they also stimulate haematopoiesis via CSF's

The class I major histocompatibility complex (MHC I) p203

- (a) class I is expressed on all *nucleated* cells of the body, and platelets
- (b) is encoded for *on chromosome 6*
- (c) has a very *high level of polymorphism*
- (d) **are encoded for by three loci designated HLA-A, HLA-B and HLA-C**
- (e) β -microglobulin *is not encoded on the MHC*

Mast cells p208

- (a) **are bone marrow derived cells**
- (b) are the primary cell involved in *type I hypersensitivity reactions*
- (c) **release PAF** and therefore must activate platelets, and are inflammatory cells
- (d) **do not phagocytose antigen**
- (e) none of the above is true *wrong*

Regarding mast cells, which statement is *false* p207

- (a) They can degranulate with exposure to morphine
- (b) They are activated with cross linking of high affinity IgE Fc receptors
- (c) They can degranulate with exposure to C5a
- (d) **Basophils are similar to mast cells, but are in much smaller concentrations, and are circulating: their functions in anaphylaxis have not been well established**
- (e) They can be activated by IL-8

An example of primary mediators (ones preformed in granules) released by Mast cells would be (p208-9)

- (a) cytokines
- (b) leukotrienes
- (c) Platelet activating factor
- (d) **Proteoglycans such as heparin, biogenic amines such as histamine, enzymes such as chymase, tryptase,**
- (e) Prostaglandin D₂

An example of secondary mediators (manufactured on signalling) released by Mast cells would be (p208-9)

- (a) histamine
- (b) **platelet activating factor and see above)**
- (c) heparin
- (d) chymase
- (e) tryptase

In systemic anaphylaxis

- (a) the severity of the response is proportional to the *level of sensitivity to the antigen*
- (b) there is widespread oedema, with striking *contraction of the respiratory bronchioles*
- (c) a previous history of some form of allergy is usually, *but not always present*
- (d) **the symptoms usually follow administration of foreign proteins**
- (e) *hives, hoarseness (larynx), vomiting, abdominal cramps, etc are a feature*

Complement membrane attack complex is activated by p210

- (a) IgE
- (b) IgA
- (c) IgD
- (d) **IgG and IgM**
- (e) none of the above

Regarding hypersensitivity reactions p211

- (a) the ***mere presence of antibody-antigen complexes in the circulation does not imply disease***
- (b) Type III hypersensitivity is generally due to reactions against ***exogenous*** antigens, but in some cases reactions to endogenous antigens cause diseases such as SLE.
- (c) Pemphigus vulgaris is an example of antibody-mediated cellular dysfunction; in this case, against desmosomes, ***which disrupt intercellular junctions in epidermis***
- (d) The ***T_H2***-type helper cell promotes the synthesis of IgE in type I hypersensitivity reactions: and it is this helper cell and the presence of increased IgE which helps promote the anaphylactic reactions
- (e) **In type III hypersensitivity, immune complexes are typically deposited in vessel walls, or extravascular sites where antigen has been deposited previously.**

In type III hypersensitivity reaction, (p213)

- (a) phase 2 (***deposition of ab-ag complexes***) begins a week after exposure to antigen
- (b) ***Large complexes formed in great antibody excess are rapidly removed by the phagocyte system, and are harmless.*** The most pathogenic antigen-antibody complexes are of small or intermediate size
- (c) wherever the complexes deposit, the tissue damage is similar: complement cascade (chemotaxis, anaphylotoxins > MAC), and n ϕ and m ϕ through activation of the Fc receptors.
- (d) phase 3 begins approximately 10 days after exposure to the antigen
- (e) **all of the above is true**

Peripheral tolerance is maintained by all of the following mechanisms except

- (a) Anergy irreversible inactivation of lymphocytes. Recognition of self without extra signals (CD28) leads to inactive lymphocytes. B cells without T cell helper activation also anergise.
- (b) **deletion of T-cells that express high affinity for self antigens** during maturation in the thymus (**central tolerance mechanism**)
- (c) clonal deletion by activation-induced cell death, via the Fas-Fas ligand system of activation induced apoptosis
- (d) Suppression by regulatory T cells (? Produce IL-10 and TGF β to inhibit activation)
- (e) Antigen sequestration (areas hidden from the immune system: testis, eye and brain, as inflammation would destroy these delicate structures)

Antigen sequestration is p225

- (a) the phagocytosis of bacteria by leukocytes
- (b) the removal of self recognising lymphocytes from the lymphocyte population
- (c) **where tissues in which antigens are located do not communicate with blood or lymph, eg brain, testis, eye: immune privileged sites**
- (d) the uptake of antigen by Peyer's patches
- (e) the presentation of antigenic products on MHC II molecules

Concerning the mechanisms of autoimmune disease p227

- (a) the actual genes associated with *most autoimmune diseases are not known, just the general chromosomal area*
- (b) normal MHC molecules are *capable of presenting self antigens*
- (c) *Many* autoimmune diseases are associated with preceding infections
- (d) molecular mimicry may play a role in the development of disease**
- (e) epitope spreading is the process by which concealed *antigens become exposed to the immune system, creating the potential for new self recognition*

Transmission of HIV p246

- (a) through needlestick injury is *approximately 0.3%, Hep B 30%*
- (b) through needlestick injury can be reduced eightfold when a patient is given antiretroviral therapy**
- (c) *intrapartum, and peripartum* is the most common mode of mother-to-infant transmission
- (d) from blood transfusion has been *virtually been* eliminated, *but is still possible* ($1:2 \times 10^7$)
- (e) it is *virtually impossible to be transmitted by insect* (thank God!)

HIV p247-8

- (a) is a single stranded RNA virus (*only rotavirus is x2 DNA*)
- (b) has two viral glycoproteins, gp120 (binds to T cell CD4, change conformation, and binds to chemokine receptor CCR5/ CXCR4) and gp41, which fuses into the cell membrane, which are vital for infection of cells**
- (c) envelope is a *lipid bilayer* from the host cell
- (d) has *tremendous* variability expressed in its genome, and it is this variability that prevents a vaccine.
- (e) invades *CD4+ cells, dendritic cells and macrophages*

Concerning HIV p249

- (a) accumulation of T tropic virus is a sign of the final rapid phase of disease progression
- (b) M-tropic strains use CCR5 chemokine receptor to infect host cells
- (c) T-tropic strains use CXCR4 chemokine receptor to infect host cells
- (d) M-tropic viruses are the dominant virus type found in newly infected individuals
- (e) all of the above is true.** M-tropic strains develop into T tropic strains owing to gp120 mutation. T-tropic viruses can infect immature T cells, and T cell precursors, seriously depleting T cell numbers. Latent infection is ceased when the T cell is mobilised and the cDNA of the virus is replicated.

Concerning HIV infection p252-3

- (a) Infected monocytes transport the virus to the brain**
- (b) The type of virus that infects the microglia is M-tropic, pointing to the fact that T cells are excluded from the immune privileged site**
- (c) Neurons are *not* infected by HIV
- (d) Macrophages are *not destroyed* by HIV virus replicating in the phagolysosome, *making them a reservoir for virus*
- (e) A and B are true**

Concerning HIV complications p256

- (a) Invasive candidiasis is ***uncommon, and only occurs if there is drug induced neutropaenia, or IDC's***
- (b) Atypical pneumonia caused by *Mycoplasma pneumoniae* is ***not*** one of the sentinel infections
- (c) **Toxoplasmosis is responsible for 50% of mass lesions in the CNS of HIV patients**
- (d) ***25-40% of infected individuals will develop a malignancy: cervical, anal (HPV), non-Hodgkin's lymphoma (most systemic type 80%), or Kaposi***
- (e) Kaposi sarcoma is ***20 times more common*** in patients who acquired the disease sexually