

## Acquired Knowledge Questions Immunology

This document include questions that aim to summarize most of the points in the immunology course and assure attaining all the needed acquired knowledge for the official exam.

Sources: Official Exams, Biocatalyst, booklet for T. Rola Dirani, Abed Al Kareem Nouredine

**1) List the signs of inflammatory reaction**

Redness, heat, pain and edema (swelling)

**2) Upon infection, damaged cells secrete certain chemicals that have local and systematic effects**

**a) Name these chemicals**

cytokines

**b) Indicate the function of these cytokines and how they lead to the signs of inflammation**

cytokines induce vasodilation leading to increase of blood flow which is the reason of redness and heat. In addition other cytokines increase vascular permeability so plasma diffuse toward infected tissue in addition to some cells that are attracted by chemokines by process called chemotaxis. The flow of plasma and cells to infected area leads to swelling.

**Note: Feeling of pain is due to of some chemicals (prostaglandin) that stimulate pain receptor in the area.**

**3) Name the mechanism by which certain leukocytes are attracted to infected area.**

chemotaxis

**4) Name the process by which monocytes leave blood vessel into infected tissue**

diapedesis

**5) Explain how macrophage reach infected tissue**

Upon infection, the infected cells, the macrophage and other leukocytes release cytokines having local and systematic effects inducing vasodilation of blood capillary in the area, which increase the blood flow. In addition, some cytokines have chemotactic effect, which attract monocytes to the site of infection by chemotaxis. The monocytes then cross the blood vessel by diapedesis where they transform into macrophage.

**6) List the phases of phagocytosis**

Adhesion, absorption, digestion

**7) What is the type of MHC where peptide of the phagocytosed antigen is presented?**

MHC II

**8) Why phagocytosis is considered as non- specific immune response?**

because the macrophages phagocytes any intruder regardless of its identity (phagocytes don't have specific receptors for specific antigens)

**9) What are the characteristics of non-specific immune response?**

No latency period, non specific

**10) There are 2 types of specific immune response**

**a) Name them**

Cell mediated immune response. Humoral mediated immune response.

**b) Indicate the effector cell / molecule of each of them.**

Cell mediated immune response: cytotoxic Tc

Humoral mediated immune response: Plasma cell / Antibodies

**11) After phagocytosis and presentation of peptide on the specific MHC, macrophage is now called..?**

APC: Antigen Presenting Cell

**12) Indicate the site of induction of immunity against antigens if the antigen is circulating through blood or through lymphatic tissues (tissue was infected)**

13) If the antigen was through blood, spleen. If the antigen was circulating through lymphatic tissue, lymph nodes.

**14) APC migrates to the nearest lymph node Or spleen**

**a) Indicate the function of lymph node**

Site of interaction between leukocytes and antigens coming through lymph

**b) Name the cell that the APC activates in lymph node.**

T4

**c) Explain the mechanism of activation/ Or Explain macrophage role in induction of specific immune response.**

The macrophage phagocytoses the non-self-antigen, it associates its peptides to HLA-II on its membrane surface.

Macrophage migrates to the closest lymph node where it becomes APC (Antigen Presenting Cell)

When antigen specific T4 arrives, it binds on the HLA-II non-self-peptide complex by double recognition with its TCR leading to its activation. This is the “clonal selection”

**d) Explain the necessity of the seven days time delay for the lymphocytes proliferation. (2008 1st)**

“7 days time delay” is necessary to induce the immune response. Macrophages phagocytose the antigens and become APC that migrate towards the lymphatic ganglia. APC bind to the lymphocytes via their specific receptors and activate them.

**15) Non specific immune response triggers specific immune response. Explain this statement**

The macrophages are the effector cells of the non-specific immune response while lymphocytes are the effectors of the specific immune response, and without macrophage, the lymphocytes can not produce antibodies.

**16) Activated T4 cells secrete certain type of cytokines**

**a) Name these molecules**

Interleukins (IL-2 and IL-4)

**b) Indicate their function**

IL-2 activates Tc that has the specific TCR for the non-self peptide and **already bound** to MHC-I non-self-peptide-complex on infected cell by double recognition through its TCR.

IL-4 activates B cells having the antibody specific to the antigen and that are **already bound to it**.

**c) Explain mechanism by which these molecules activate their target cells (activation of B or T lymphocytes).**

IL-2 secreted by T4 cells activates only T8 that has the specific TCR and already bound to the peptide MHC-1 complex by double recognition. This is clonal selection. This leads to multiplication of the activated T8 obtaining slightly differentiated memory cells and differentiation leading to having cytotoxic Tc

IL-4 secreted by T4 cells activates only Bc that has the specific immunoglobulin to the antigen and already bound to it. This is clonal selection. This leads to multiplication of the activated B cells obtaining slightly differentiated memory cells and differentiation leading to having plasma cells.

**17) Explain how Plasma cells (plasmocytes) are cells adapted for secretion of antibodies. (2017 1<sup>st</sup>)**

Plasma cells have a well-developed cytoplasm rich in rough endoplasmic reticulum involved in protein synthesis and since antibodies are proteins, therefore the plasma cell is a cell adapted to the secretion of antibodies

**18) Explain the mechanism of antibodies against toxins leading to their neutralization**

Toxins attach the target cells by binding to membrane receptors. When the antibodies bind to the toxins they cover their attachment sites on the target cells and prevent the toxin from acting. This is neutralization.

**19) Explain the 2 ways how antibodies may eliminate the antigen.**

**By opsonization:**

After binding of the antibodies using their Ag binding sites of variable region to the epitope of the antigen, the antibody can bind to membrane receptors of macrophage through its constant region making a molecular bridge between antigen and the phagocyte that facilitates adhesion. **This is opsonization.** The macrophage phagocytes the antibody with antigen bound to it in order to eliminate them.

**Activation of complement:**

Upon the binding of antibodies by their variable region on the bacterial antigens, a complement protein C1, may bind to constant region and become activated. This activation leads to complement cascade activation from C1 to C9 that forms at the end a membrane attack complex (MAC) on surface of the bacteria that perforate the membrane of this cell destroying it.

**20) Explain the role of antibodies in defense against viral infection.**

Viruses enter their target cell through the binding of their surface molecules to a specific receptor expressed by the membrane of the target cell. By binding to these surface molecules. The antibodies can prevent the entry of the virus into the host cell and thus neutralize its activity.

**21) Specific immune response facilitates the function of non- specific immune response. Explain this statement.**

Upon binding of antibodies to their specific antigen (that is a specific immune response). The antibody can bind to membrane receptor of macrophage through its constant region facilitating adhesion of phagocyte and perform phagocytosis that is non-specific. So specific immune response facilitate the function of non-specific immune response

**22) Indicate the roles and the moments where macrophages intervene in the specific immune response (2015 1<sup>st</sup>).**

At the beginning of the specific immune response, macrophages act as antigen presenting cells which induce the specific immune response.

At the end of the specific humoral immune response, they perform phagocytosis of the immune complex in order to eliminate antigens

**23) Upon antigen infection, immune complexes appear after certain time but then disappear. Explain (2008 1<sup>st</sup>)**

The appearance of immune complexes is due to the neutralization of the antigen by the antibodies secreted by plasmocytes. The disappearance of these complexes is due to the opsonisation and phagocytosis carried out by macrophages.

**24) Explain how secreted antibodies contribute to the destruction of a virus. (refer to 2011 1<sup>st</sup>)**

The specific antibodies neutralize their corresponding antigens of the flu virus by binding to them through their specific antigenic binding sites forming immune complexes. Thus the antibodies become able to bind through their constant part on macrophages that phagocyte the whole immune complexes thus destroying the virus (opsonization).

**25) Indicate the type of foreign antigens that triggers cell-mediated specific immune response, and give an example**

Intracellular antigens such as viruses.

**26) Justify the necessity of cell mediated immune response to eliminate virus from the body.**

Viruses are intracellular pathogens can only multiply within the host cell, elimination of the virus from the body requires destruction of cells infected by the virus which is performed by T<sub>C</sub> lymphocytes, effectors of the specific cell mediated immune response

**27) Explain why nude mouse (lack thymus) lacks humoral and cell mediated immune response**

In the absence of thymus, T<sub>L</sub> cannot be mature so there is absence of T<sub>C</sub> leading to no cell-mediated specific immune response. Moreover, there is no mature T<sub>4</sub> which cannot be activated to secrete interleukin 4 so cannot activate B-lymphocytes. Therefore, humoral mediated immune response will not be activated

**28) Compare between memory cells and effector cells (differentiated) regarding ability of proliferation and life span**

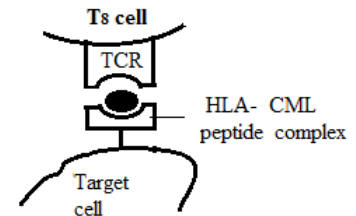
Memory cells have long life span and can proliferate while differentiated cells can not proliferate and have short life span.

**29) Explain how the anti-CD4 antibodies intervene in accepting grafts. (2008 2nd)**

They fix to TL4 receptors to block them. The blocked TL4 are not activated and do not proliferate nor do they differentiate into cells that secrete interleukins 2. Thus, the TL8 are not activated and the graft is successful. Anti-CD4 antibodies can be used as immunosuppressor drugs during grafting.

**30) Draw a scheme showing the molecules involved in the recognition taking place between the activated T8 cell and the target cell.**

Title: Scheme of the recognition site between T8 cells and the target cell



**31) Indicate the conditions necessary for T4 proliferation (refer to 2017 2<sup>nd</sup>) + additional information.**

T4 recognize the non-self-peptide MHC II complex by double recognition (note: same as for T8 cells by the way) for that:

cooperation between T4 and macrophage is necessary for T4 proliferation

T4 and macrophage must be from the same strain. (2017 2<sup>nd</sup>)

T4 must have the TCR specific for non-self-peptide MHC II complex presented on macrophage.

**32) Specify the consequence of the absence of the macrophages on the specific immune responses. (2017 2<sup>nd</sup>)**

The induction of specific immune response ceases because the activation of T4 lymphocytes necessitate its binding to APC. So in the absence of activated T4 lymphocytes, no more secretion of interleukin 2 takes place which is responsible for launching the specific cell mediated immune response. Moreover, no interleukin 4 secretion takes place which is responsible for launching of the specific humoral immune response.

**33) Explain the mechanism that leads to the destruction of target cells (infected cells) by Tc lymphocytes (2010 2<sup>nd</sup>)**

The T<sub>C</sub> lymphocytes recognize and bind, by its TCR, to target cells expressing the modified self: self MHC I carrying a non-self-peptide of the antigen that is at the origin of their activation. They will then release, by exocytosis, perforin molecules forming hollow polyperforin channels through the cell membrane, and then they release granzymes molecules that penetrate the target cell through these channels leading to DNA degradation and to the target cell destruction.

**34) Explain how the cells: macrophage, T4 and T8 intervene in cell lysis (2007 1<sup>st</sup>)**

The macrophages digest the free virus, recognized as non-self, and transform them into peptides and present them on HLA molecules of class II. These macrophages are thus, antigen presenting cells (APC). These latter migrate towards the lymphatic ganglia where they activate the LT4 that has TCR specific for the peptide expressed on APC that secrete IL-2. IL-2 activates the specific LT8 (LTc), which adheres to the membrane of the target cell and binds to MHC-I non self peptide complex by its TCR by double recognition and releases perforin that forms polyperforin channel and granzymes that enters through polyperforin channel and degrades the DNA leading to cell lysis.

**35) Indicate the characteristics of 2ary immune response:**

More amplified, faster, long lasting.

**36) Note: To prove the following characteristics:**

**More amplified:** compare the number of antibodies produced or number of specific T8 cells between 1<sup>st</sup> contact and second contact.

**Faster:** Compare the time needed to trigger the immune response (when the antibodies increase or T8 cells)

**Long lasting:** Compare the final amount shown in graph where upon 2ary contact the amount of antibodies would be much higher than 1<sup>st</sup> contact which reaches 0 after certain time

**37) Explain the origin of characteristics of the 2ary immune response/ or the results in experiments shows the elimination of antigen upon second contact is faster (in less time) explain the results.**

During the secondary immune response there are memory cells that are more numerous and more differentiated than naïve cells (B or T cells depending on your exercise) sensitized during the primary immune response. These memory cells are able to recognize the antigen faster, proliferate, and differentiate to a larger number of plasma cells that secrete higher concentration of antibodies persist for a longer time (or activated T8 depending on the exercise)

**38) Indicate how does vaccine protect against antigen? (2013 2<sup>nd</sup>)**

Vaccines immunize the organism against a specific antigen by inducing a durable immunological memory.

**39) Explain the importance of vaccine (2016 1<sup>st</sup>)**

Vaccine ensures the first contact with this antigen and triggers immunological memory. Consequently, the body, after a second contact, develops a secondary response which is more amplified, more rapid and more durable against this antigen

**40) Given that infection with a certain virus causes cervical cancer. Explain how vaccination against this virus protects against cervical cancer. (2015 2<sup>nd</sup>)**

The antibodies produced due to vaccination neutralize the viruses before they bind to the membrane receptors of the target cells of the cervix and inhibit the viruses from infecting them.

The viruses are thus eliminated (the formed immune complexes will be phagocyted by macrophages).

**41) Distinguish between vaccination and serotherapy (2015 1<sup>st</sup>). Refer to this table for information but write your answer as text by comparing each criteria**

	<b>Vaccination</b>	<b>Serotherapy</b>
<b>Nature of the injection</b>	Killed or attenuated antigen	Serum contacting antibodies
<b>Origin of antibodies</b>	Endogenous	Exogenous
<b>Latency period</b>	2 weeks	null
<b>Duration of protection</b>	Several years ( <b>more durable</b> )	Two weeks ( <b>short</b> )
<b>Nature of immunity acquired</b>	Active	Passive

**42) You might be asked to specify the importance of a certain step of experimental protocol. Usually, this step is necessary to assure presence of only one variable in order to reach a proper conclusion.**

**Strategy of thinking: Ask yourself what would happen if this step wasn't included or different?**

**Examples as following:**

**Example 1: Refer to 2012 2<sup>nd</sup> :**

In the framework of determining the conditions of the production of antibodies during the immune response, we perform a series of experiments on mice of the same strain.

Experiment Mice are subjected to the ablation of the thymus followed by irradiation that destroys all cells of the immune system. These mice are then divided into 4 lots and treated as shown in document 1. (injection with different types of lymphocytes)

**a) Specify the aim of destroying the cells of the immune system before starting the experiment.**

This will ensure that the immune response triggered by the mice is due **only** to the injected cells and not due to cells presented already in the body.

**b) Specify why the injected lymphocytes should be from the same strain.**

T lymphocytes by their TCR recognize peptides within MHC complex by double recognition (recognize the non self peptide and recognize the MHC complex.) So if the T lymphocytes were from other strain they would have different MHC which can't be recognized by T lymphocytes.

**Example 2:**

To study the immune response against tumors, several experiments were done started by graft of tumor to a mouse from the same strain (same MHC).

**Explain why the tumor transplant should be from mouse having same MHC (or same strain)**

If the tumor is of different MHC, it can be rejected due to the different MHC marker but not because of its tumor character. On the other hand, a grafted tumor from a different MHC mouse presents for the experiment two variable factors that do not allow us to draw the conclusion properly for the problem.

**43) Name the 2 types of immunodeficiency**

Congenital immunodeficiency, acquired immunodeficiency

**Explain the outcomes in patient having mutation in gene coding for IL-2 receptor**

T8 cells are activated by IL-2 secreted by T4. Thus, T8 cells of patient suffering from mutation in IL-2 receptor can not be activated. And so this patient lacks cell mediated immune response and is not able to fight intracellular pathogens thus having immunodeficiency in cell mediated immune response.

**44) Name the virus leading to immunodeficiency**

HIV

**45) What is his target cell(s)?**

HIV target cells are cells having CD4 receptors (mainly T4 and macrophages)

**46) Explain the development of opportunistic diseases by AIDS patient.**

Amount of T4 cells should be greater than (usually  $200/\text{mm}^3$ ) (but refer to document if presented) is essential to prevent

T4 cells are required for activation of T8 cells by secreting Interleukin 2 and B cells by Interleukin 4 which becomes activated cytotoxic Tc and plasma cells respectively. These are the effector cells for cell mediated and humoral mediated immune response.

An AIDS patient having T4 cells lower than  $200/\text{mm}^3$  (or refer to document in exercise for number) which is below threshold of immunity are not sufficient for activation of T8 and B cells, consequently lacks cell mediated and humoral mediated immune response which allows the development of opportunistic diseases.

**47) A diabetic rat is treated with cyclosporine (immunosuppressive substance), it becomes healthy.**

**Explain**

This means that the disease is due to the immunity since once attenuated the disease is cured. Therefore, the diabetes of this rat is due to an autoimmune disease.

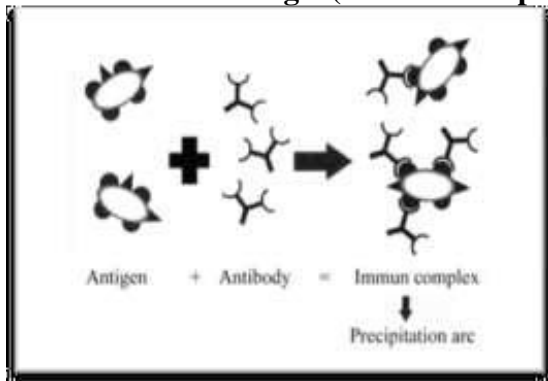
**48) An arthritic rat, is subjected to thymus ablation, after a while it becomes healthy. Explain**

This result means that the disease is due to the thymus (immunity), because without thymus there is no mature T lymphocytes and the disease disappears. Therefore, the disease is an autoimmune disease.

**49) Injection of antibodies of affected individual into a normal one leads to same disease (same symptoms) that disappear after a while. Explain**

This result means that the disease is due to the antibodies which transmitted the disease to the other individual and the disease disappeared after the amount of injected of antibodies decline. Therefore the disease is due to an autoimmune disease

**50) Schematize a diagram showing the mechanism that leads to the formation of the precipitation arc in immunodiffusion in gel (immune complex). (2010 1<sup>st</sup>)**



**51) Name the technique used for detection of HIV virus.**

ELISA

**52) Name the technique used for detection of specific protein, receptor..?**

Immunofluorescence