

إبراهيم البيشي :DONE BY ماجد العسبلي عبدالاله ابو خلف

> Thank you for checking our work, Good luck! If you have any suggestions or alterations contact us! Email <u>Immunology435@gmail.com</u>

Major Histocompatibility Complex and Transplantation

- Human Leukocyte Antigens (HLA): Are genes that encode for the MHC protein, and are found on the short arm of chromosome 6.
- For a transplantation procedure to be successful, there has to Be an MHC Matching between the donor and the acceptor.
- Each individual has two "haplotypes" i.e, two sets of these genes one paternal and one maternal
- MHC Class II Has a stronger effect on tissue rejection than class I.

	MHC Class 1	MHC Class II
Gen Products	HLA-A,-B,&-C	HLA-DP,DQ
Cells Distribution	all nucleated cells	antigen presenting cells
		(macrophages, B cells, dendritic cells and Langerhans cells)
Recognized By	Cytotoxic T cell (CD8+)	Helper T cell (CD4+)



Minor HLA genes and Transplantation

- <u>Minor HLA genes unknown</u>
 - They mount a weak immune response
 - Play role in chronic rejection of a graft
 - -There are no laboratory tests to detect minor antigens

Types of Transplantation



Immune responses to transplantation (Rejection):

- Major Barrier to transplantation is the immune response
 - T Cells: Has a major role (1st & 2nd set reactions). CD4 cells are more effective than CD8 cells.
 - B cells: May play a role (Antibodies)
 - Classic adaptive/acquired immune response
 - Memory: Activated as a type of a secondary response if the same graft is transplanted again (2nd set reaction).
 - \circ Specificity

Type of rejection	Onset	Features
Hyperacute rejection:	very quick	<i><u>caused by</u>.</i> preexisting host serum antibodies specific for antigens of the graft
Acute rejection:	about 10 days (cell mediated)	
Chronic rejection:	months-years (both) (cell mediated &Ab mediated)	- <u>Main pathologic finding</u> :is atherosclerosis of the vascular endothelium
		- <u>Main cause</u> :is not known. <u>Could be due to:</u>
		 Minor histo-compatibility antigen miss match
Graft-versus-Host (GVH) Reaction		<u>Definition:</u> Donor's Tc cells play a major role in
		destroying the recipient's cells
		<u>Cause:</u>
		grafted immunocompetent T cells proliferate in the irradiated immunocompromised host and reject cells with foreign proteins resulting in sever organ dysfunction
		<u>Occurs in:</u>
		about two thirds of bone marrow transplants
		<u>Symptoms are:</u>
		maculopapular rash, jaundice, hepatosplenomegaly and diarrhea
		<u>Prognosis:</u>
		GVH reactions usually end in infections and death

Tissue Matching:

Cornea	Kidney	Bone marrow
Immunosuppression not required	ABO and HLA matching useful Immunosuppression usually required	ABO and HLA matching req

Laboratory testing for HLA matching:

HLA\Tissue typing: To determine if donor and recipient's HLA match or not before transplantation

Methods:

- 1) DNA sequencing by PCR.
- 2) Serologic Assay.

3) Mixed Lymphocyte Reaction (MLR).

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4) Cross-matching.

Immunosuppression therapy:

_Divided into:

- 1) <u>General immunosuppression.</u>
 - A. Mitotic inhibitor: azathioprine (pre & post)
 - B. Corticosteroids
 - C. Cyclosporine
 - D. Total lymphoid irradiation
- 2) Specific Immuno-suppression.
 - i) Monoclonal antibodies against T cell components or cytokines
 - ii) Agents blocking co-stimulatory signal (Anergy)

_Downsides

- Must be maintained for life
- Toxicity
- Susceptibility to infections
- Susceptibility to tumors



1-genes for HLA protein are clustered in (MHC complex) located on the short arm of which of the following:	2-which of the following play a primary role in rejection reaction:	3-in Graft Versushost (GVH) reaction which of the following play a primary role:
<u>A-chromosome 6.</u> B-chromosome 17. C-chromosome 8.	A-B cells. B-adaptive response. <u>C-T cells.</u>	A-donor B cells. B-recipient Tc cells. <u>C-donor Tc cells.</u>
 4-which of the following is general immunosuppressive therapy: A-antibody against T cell component. <u>B-total lymphoid irradiation.</u> C-blocking co stimulatory signals 	5-which of the following is a symptom in Graft Versus Host reaction: A- maculopapular rash. B- hepatosplenomegaly. <u>C- all above.</u>	6- Xenogeneic grafting means donor and recipient are different species ? A- <u>True.</u> B-False.
7- Chronic rejection occurs with in ? A- hours. <u>B- months to years.</u> C- days.	 8- GVF reactions occur in : <u>A- two thirds of bone marrow</u> <u>transplants.</u> B- one fifth of bone marrow transplants. C- does not occur in bone marrow transplants. 	 9- Syngeneic grafting means that: A- donor and recipient are the same. B- donor and recipient are genetically identical. C- donor and recipient are same species.
10- The success of tissue and organ transplantation depends upon the donor's and recipient's ? <u>A- HLA.</u> B- Blood type. C- age.	1-A. 2-C. 3-C. 4-B. 6-A. 7-B. 8-A. 9-B.	5-C. 10-A.