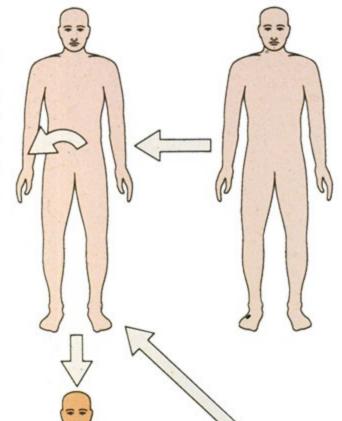
Immunology of transplantation

Types of transplantation

- Autotransplantation –within one organism
- Allotransplantation- between one species
- Xenotransplantation- between two different species

autograft

from one part of the body to another e.g. trunk to arm



isograft

between genetically identical individuals e.g. monozygotic twins,or within an inbred strain

allograft

between different members of the same species e.g. Mr Smith to Mr Jones

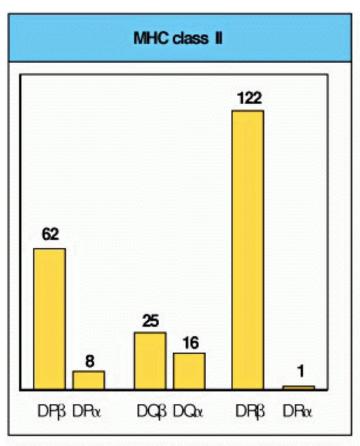


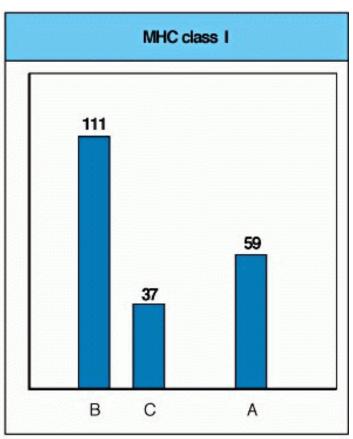
between members of different species e.g. monkey to man

Success rate of transplantation in humans

Tissue transplanted	5-year graft survival*	No. of grafts in USA (1999)
Kidney	80-90%	13,429 (12,483)
Liver	40-50%	4698
Heart	70%	2234 (2185)
Lung	30-40%	934 (885)
Comea	~70%	~40,000†
Bone marrow	80%	23,500‡

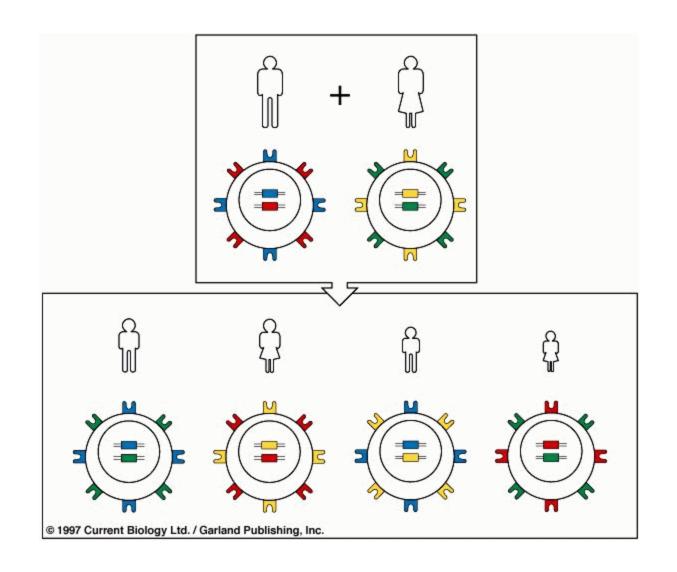
Polymorphism of human MHC antigens



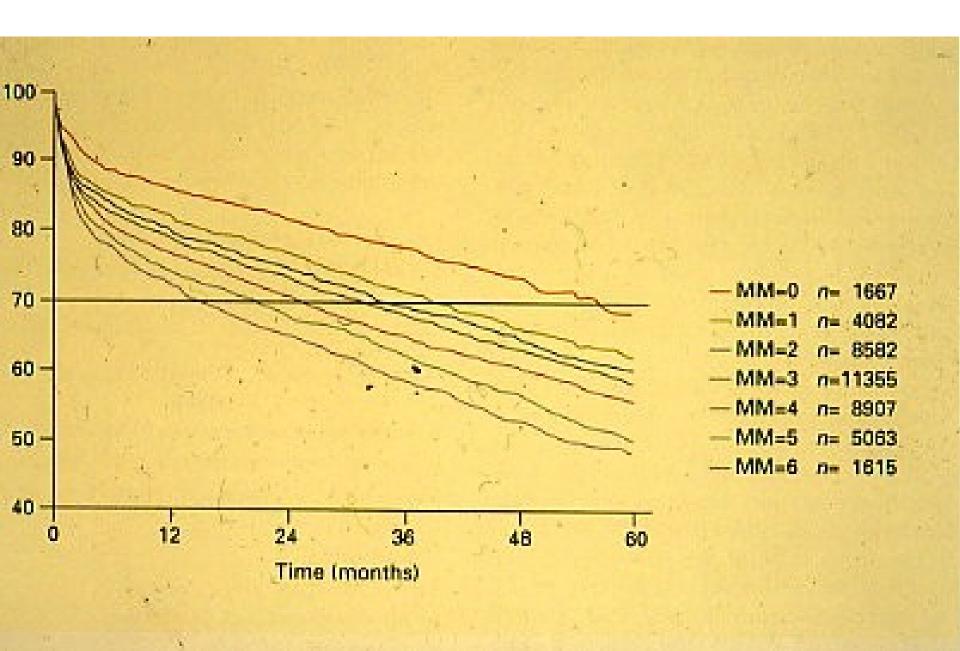


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Co-dominant expression of HLA genes



Effect of HLA-identity on kidney graft survival



Types of graft rejection

- Hyperacute minutes to few hours after transplantation. Caused by pre-formed antibodies against HLA antigens of the donor. Irreversible.
- <u>Acute</u> -several days to months after transplantation. Mainly T-cell mediated. Usually reversible by aggressive immunosuppression.
- Chronic years after transplantation.
 Continuous decrease in graft function.
 Irreversible. Mechanism unknown.

The most frequent types of organ transplantation

- Heart
- Kidney
- Liver
- Lungs
- Pancreas
- Cornea

Heamatopoietic stem cells transplantation

- Indications: malignancies, bone marrow failure, primary immunodeficiencies.
- "Whole" bone marrow or separated CD34+ cells can be used.
- The most significant complication: graftversus host reaction.
- Optimal HLA-matched donor is required.

Graft-versus host reaction

- Immunological reaction of transplanted T-cells against recipients (HLA) antigens.
- Skin, liver, intestine predominantly affected.
- Milder forms can be treated by immunosuppression, severe forms may be fatal.
- Can be induced by transfusion of non-irradiated blood to immunodeficient patients (primary immunodeficiencies, leukemia...).

Systemic Immunosuppression

- High-dose steroids
- Purine antagonists: Azathioprin
- Alkylating agents: Cyclophosphamide
- Anti-pholates: Methotrexate
- Calcineurin antagonists: Cyclosporine A, Rapamycin, Tacrolymus
- Block of purins synthesis: Mycophenolate
- Antilymphocytic serum
- Monoclonal antobodies: anti CD3, anti CD25...