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Principle of Transplantation
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LEARNING OBJECTIVES

- The immunological basis of allograft rejection
- The principles of immunosuppressive therapy
- The side effects of nonspecific immunosuppression
- The major issues concerning organ donation

- The main indications for organ transplantation
- The surgical principles of organ implantation
- The causes of graft dysfunction
- The likely outcomes after transplantation

Definitions

- Allograft: an organ or tissue transplanted from one individual to another
- Xenograft: a graft performed between different species
- Orthotopic graft: a graft placed in its normal anatomical site
- Heterotopic graft: a graft placed in a site different from that where the organ is normally located
- Alloantigen: transplant antigen
- Alloantibody: transplant antibody
- HLA: human leucocyte antigen, the main trigger of graft rejection

GRAFT REJECTION

ABO blood group antigens

Permissible transplants are:

- group O donor to group O, A, B or AB recipient;
- group A donor to group A or AB recipient;
- group B donor to group B or AB recipient;
- group AB donor to group AB recipient.

There is no need to take account of Rhesus antigen compatibility.

HLA antigens

- Are the most common cause of graft rejection
- Their physiological function is to act as antigen recognition units
- Are highly polymorphic (amino acid sequence differs widely between individuals)
- HLA-A, -B (class I) and -DR (class II) are the most important in organ transplantation
- Anti-HLA antibodies may cause hyperacute rejection

Types of graft rejection

Hyperacute

- Immediate graft destruction due to ABO or pre-formed anti-HLA antibodies.
- Characterised by intravascular thrombosis

Acute

- Occurs during the first 6 months
- T-cell dependent, characterised by mononuclear cell infiltration
- Usually reversible

Chronic

- Occurs after the first 6 months
- Most common cause of graft failure
- Non-immune factors may contribute to pathogenesis
- Characterised by myo-intimal proliferation in graft arteries leading to ischaemia and fibrosis

Staging of transplantation in surgical practice

- Urgent Transplantation .
- Planned Transplantation .

Requirements

- Recognized Transplantation Center.
- Bank of Information.
- Transplantation Team :
 - 1. Immunologist.
 - 2. Hematologist.
 - 3. Transplant Surgeon: Gen/Vascular/special e experience.
 - 4. Double operating theater.
 - 5. Trained operating nursing staff.
- Donor.
- Recipient patient.
- Postoperative care unit.
- Post transplantation outpatients clinic.
- IMMUNOSUPPRESSIVE THERAPY.

Overcoming the shortage of organs for transplantation

- Maximizing heart-beating deceased donation
- Use of marginal heart-beating deceased donors
- Use of non-heart-beating donors
- Use of split-liver transplantation
- Increased living donor kidney (and liver) transplantation

Clinical testing for brainstem death

- Absence of cranial nerve reflexes.
- Absence of motor response.
- Absence of spontaneous respiration.

Absence of cranial nerve reflexes

Two physicians should be involved (Neurosurgeon & Intensive care specialist).

The treating physician should not be included.

- Pupillary reflex
- Corneal reflex
- Pharyngeal (gag) and tracheal (cough) reflex
- Oculovestibular (caloric) reflex

Absence of motor response

- The absence of a motor response to painful stimuli applied to the head/face and the absence of a motor response within the cranial nerve distribution to adequate stimulation of any somatic area is an indicator of brainstem death
- The presence of spinal reflexes does not preclude brainstem death.

Absence of spontaneous respiration

- After pre-ventilation with 100% oxygen for at least 5 min, the patient is disconnected from the ventilator for 10 min to confirm absence of respiratory effort, during which time the arterial Pco2 level should be > 8 kPa (60 mmHg) to ensure adequate respiratory stimulation.
- To prevent hypoxia during the apnoeic period, oxygen (6 | min-1) is delivered via an endotracheal catheter

Factors determining organ function after transplantation

- Donor characteristics.
- Procurement-related factors.
- Recipient-related factors.

Donor characteristics

- Extremes of age.
- Presence of pre-existing disease in the transplanted organ.
- Hemodynamic and metabolic instability.

Procurement-related factors

- Warm ischemic time.
- Type of preservation solution.
- Cold ischemic time Recipient-related factors.

Recipient-related factors

- Technical factors relating to implantation
- Hemodynamic and metabolic stability
- Immunological factors
- Presence of drugs that impair transplant function