### Transplantation

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### Transplantation

- Transfer of living tissue
- Cells: stem cells, blood cells platelets, ...
- Tissue: blood, bone marrow, skin, bone, cartilage, cornea, vessel, heart valve, fat tissue
- Organ: kidney, heart, lung, liver, pancreas, small intestine, uterus, spleen, ovary
- Body parts: hand/upper limb, face

#### Types of transplantations

- The four major types of grafts are:
  - Autografts graft transplanted from one site on the body to another in the same person (skin, vessel, blood, ovary, heart valve)
  - Isografts grafts between identical twins (1. successfull transplantation kidney 1954)
  - Allografts transplants between individuals that are not identical twins, but belong to same species (human-human), most common
  - Xenografts grafts taken from another animal species (skin, heart valves; pighuman)

### Types of transplantations - localization

- Heterotopic: to a different position (kidney into pelvic region, pancreatic islets into mesenterium), the recipient's own organ remains in its place
- Orthotopic: to the same anatomic position
  - Necessity of prior removal of the recipient's own organ (heart, lung, liver) explantation
  - Implantation of the donor organ (graft)

### Types of transplantations

- Combined organ transplantations possible, many combinations
  - Heart + kidney
  - Heart + lungs
  - Heart + liver + kidney
  - Liver + kidney + pancreas + intestine + spleen

### Organ transplantation

- Treatment of end-stage organ failure
- Temporary/auxiliar functional replacement (uterus, liver)
- Waiting list normal /urgent
- Cadaver donor most common
- Living donor for kidney, part of the liver, skin, ...
- Selection of most suitable donor and recipient <u>ABO-system most important</u>
- Ultimate goal: most possible immunologic tolerance

#### Organ transplantation problems

- Shortage of available and suitable organs
- Preservation of donor organs (time outside the blood circulation)
- Surgical techniques of transplantation
- Immunosuppression therapy to prevent rejection
- Diagnosis of possible rejection, infections, surgical complications

### Posttransplantational complications

- Ischemic injury (stop of blood flow), reperfusion injury (after implantation)
- Rejection, GVHD graft-versus-host-disease (bone marrow transpl.)
- Immunosuppression complications (opportunistic infections, neoplasia – 100x increased incidence; drug cytotoxicity)
- Other complications (surgical, original disease recurrence)
- Organ retransplantation sometimes possible

#### Organ transplantation problems

- Rejection: complex immunologic process, cellular and humoral reaction
- **Factors** genetic diversity, type of tissue (vascularisation, number of antigen presenting cells), host immune system activity (immunosuppression), graft condition
- Rejection in reaction on presence (+ demasking grade) of foreign antigenes
- Hyperacute, acute, chronic rejection

#### Opportunistic infections

Risk due to acquired immunodeficiency

- Viral activation of opportunistic microorganisms cytomegalovirus CMV, Ebstein-Barr virus EBV
- Mycotic ubiquitous fungi (aspergillus)
- Bacterial TB, common bacteria
- Parasitic toxoplasma

- Postoperatively:
  - Long recovery period
  - Side effects of long-term immunosuppression
    - Diabetes mellitus, accelerated hyperlipidemia
  - Lifelong changes drug compliance, diet changes, ...

- Before transplantation: long term poor health, severe deconditioning, exercise intolerance
- Pretransplantation activity necessary to maintain function
- Training to maintain/increase muscle strength before adverse effects of steroid therapy

- Immediate start of physical therapy after transplantation necessary
- Various training programs aerobic, muscle endurance, resistive training
- Improved quality of life
- Persistent limitations common (decrease in workload, earlier onset of anaerobic threshold, lower exercise capacity)
- Denervation of transplanted organs loss of autonomic response (heart, kidney); no problems in liver
- Lungs delay in bronchodilatation longer warm-up period necessary

- Musculoskeletal effects: osteoporosis, vertebral fractures, myopathies
- Neurotoxic reactions: tremor, paresthesia
- GIT problems
- Decreased wound healing

# Hematopoietic cell transplantation – implications for the therapist

- Treatment of hematologic neoplasias (leukemia, lymphoma, myeloma)
- Non-neoplastic blood disorders (bone marrow failure aplastic anemia, inborn severe combined immunodeficiency)

# Hematopoietic cell transplantation – implications for the therapist

- Short and/or long-term complications 30% lower life expectancy
- Immunodeficiency
- Loss of immune memory (vaccines, after infections, ...)
- Bone marrow failure
- Sterility
- Neurocognitive impairment
- Cardiopulmonary toxicity
- Graft-versus-host disease

### Hematopoietic cell transplantation – implications for the therapist

- Assessment of past life/medical history,
- Prior level of function/exercise/activities
- Assessment of general/specific condition
- Knowledge of specific medical regimen

- Risk for imobility, pneumonia, pressure ulcers, muscle weakness
- Skin care
- Oral mucositis

### GVHD, implications for the therapist

- GVH disease occurs in any situation in which immunologically competent cells or their precursors are transplanted into immunologically crippled recipients, and the transferred cells recognize alloantigens in the host
- May be fatal
- Most important complication of hematopoietic cell transplantation

#### Graft-versus-host disease

- In most patient with bone marrow transplantation, possible in organs with higher amount of lymph. tissue – intestine, liver (immunologic competent T cells + precursors → in immunodeficient host)
- HLA typization necessary
- hyperacute 7-14 d., fever, generalized erythrodermia
- acute –skin rash, mucosal ulceration, liver cholestatic lesions, thrombocytopenia, anaemia
- chronic chron. lichenoid lesions + atrophy of skin, mucosa, bronchiolitis obliterans, chron. hepatitis,...

### GVHD





#### Graft-versus-host disease

- Possible signs observed by the therapist:
  - Progressive dyspnea
  - Heart palpitations
  - Chest pain
  - Increasing fatigue
- Neuromusculoskeletal problems
  - Generalized polyneuropathy
  - Muscle wasting
  - Joint pain + stiffness, contractures (in chronic GVHD)
  - Deep tendon reflexes changes