

Organs and Organ Systems

ChemEng 590B: Tissue Engineering

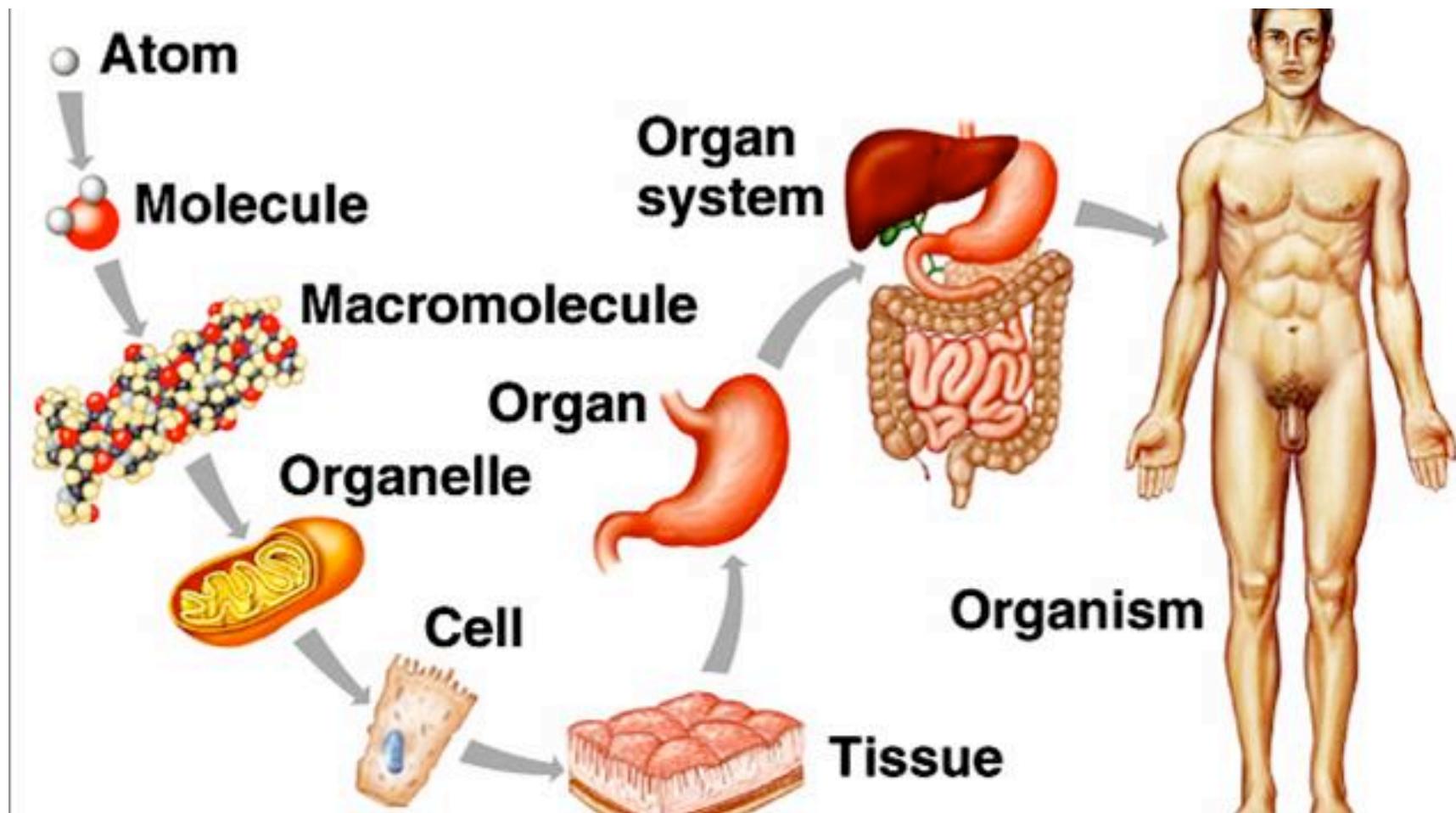
Lecture 3

January 29th, 2013

Definitions, top down

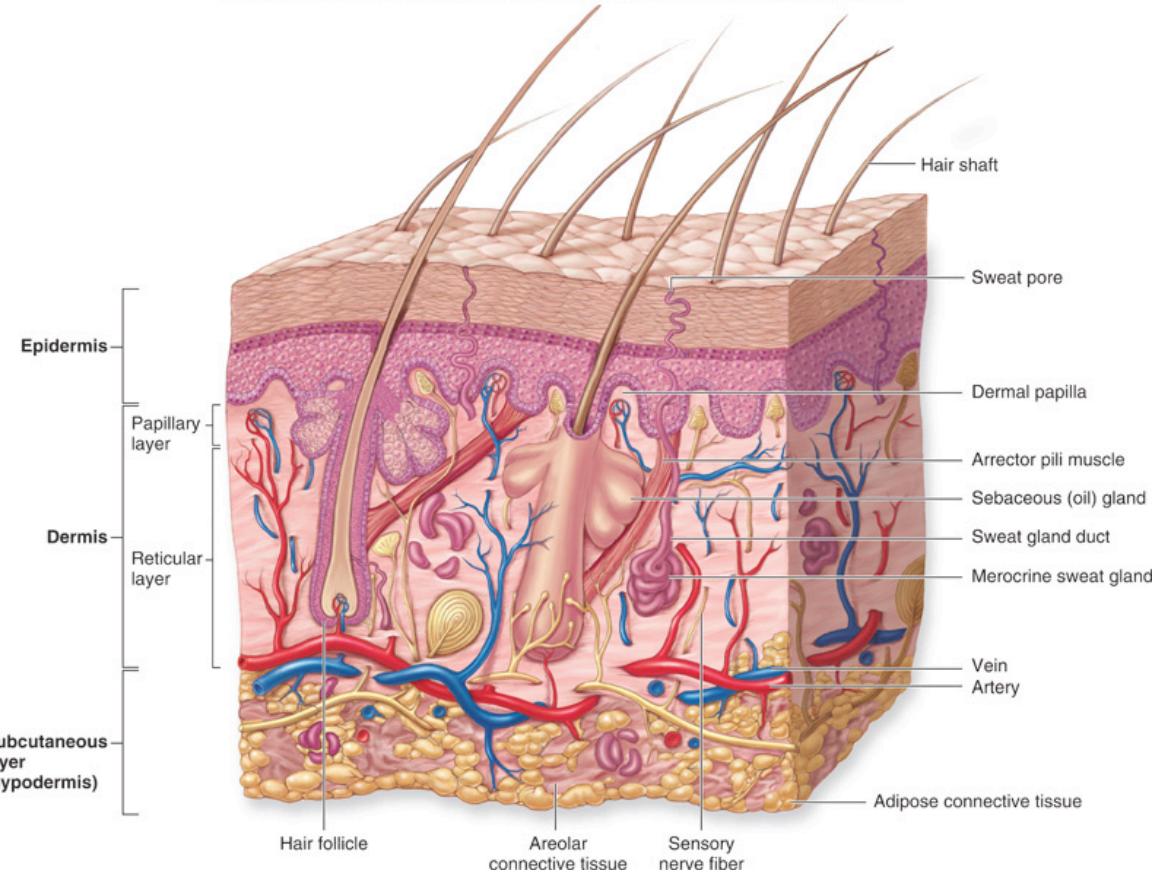
- Organism: any living thing considered as a whole.
 - E.g. Human. Bacterium.
- Organ system: group of organs that have a common function or set of functions and are therefore viewed as a unit. Humans have 11 major organ systems
 - E.g. respiratory, digestive.
- Organ: composed of two or more tissue types that perform one or more common functions.
 - E.g. bladder, heart, skin
- Tissue: group of similar cells and the materials surrounding them.
 - E.g. epithelial, connective, muscle
- Cell: the basic unit of all living things.
- Organelles: combined of molecules, and make up the small structures that make up cells.

Levels of Organization



Integumentary System

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Provides protection, regulates temperature, prevents water loss, and produces Vitamin D precursors

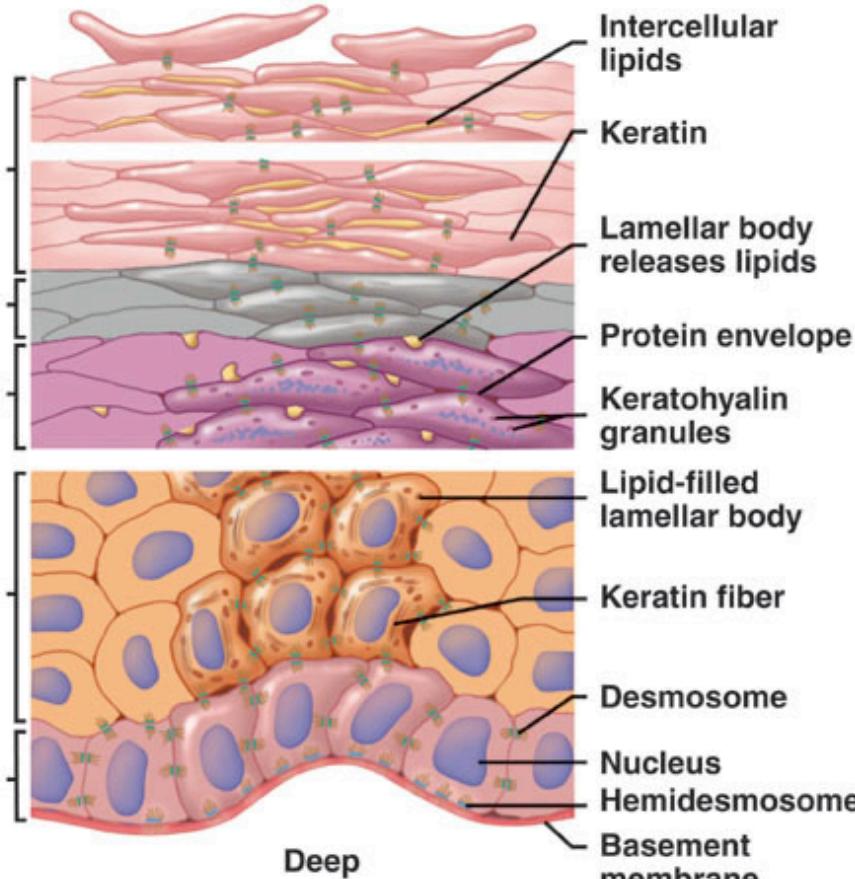
Consists of skin, hair, nails, and sweat glands

Epidermal Layers of the Skin

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Superficial

5. **Stratum corneum**
Dead cells with a hard protein envelope; the cells contain keratin and are surrounded by lipids.
4. **Stratum lucidum**
Dead cells containing dispersed keratohyalin.
3. **Stratum granulosum**
Keratohyalin and a hard protein envelope form; lamellar bodies release lipids; cells die.
2. **Stratum spinosum**
Keratin fibers and lamellar bodies accumulate.
1. **Stratum basale**
Cells divide by mitosis and some of the newly formed cells become the cells of the more superficial strata.



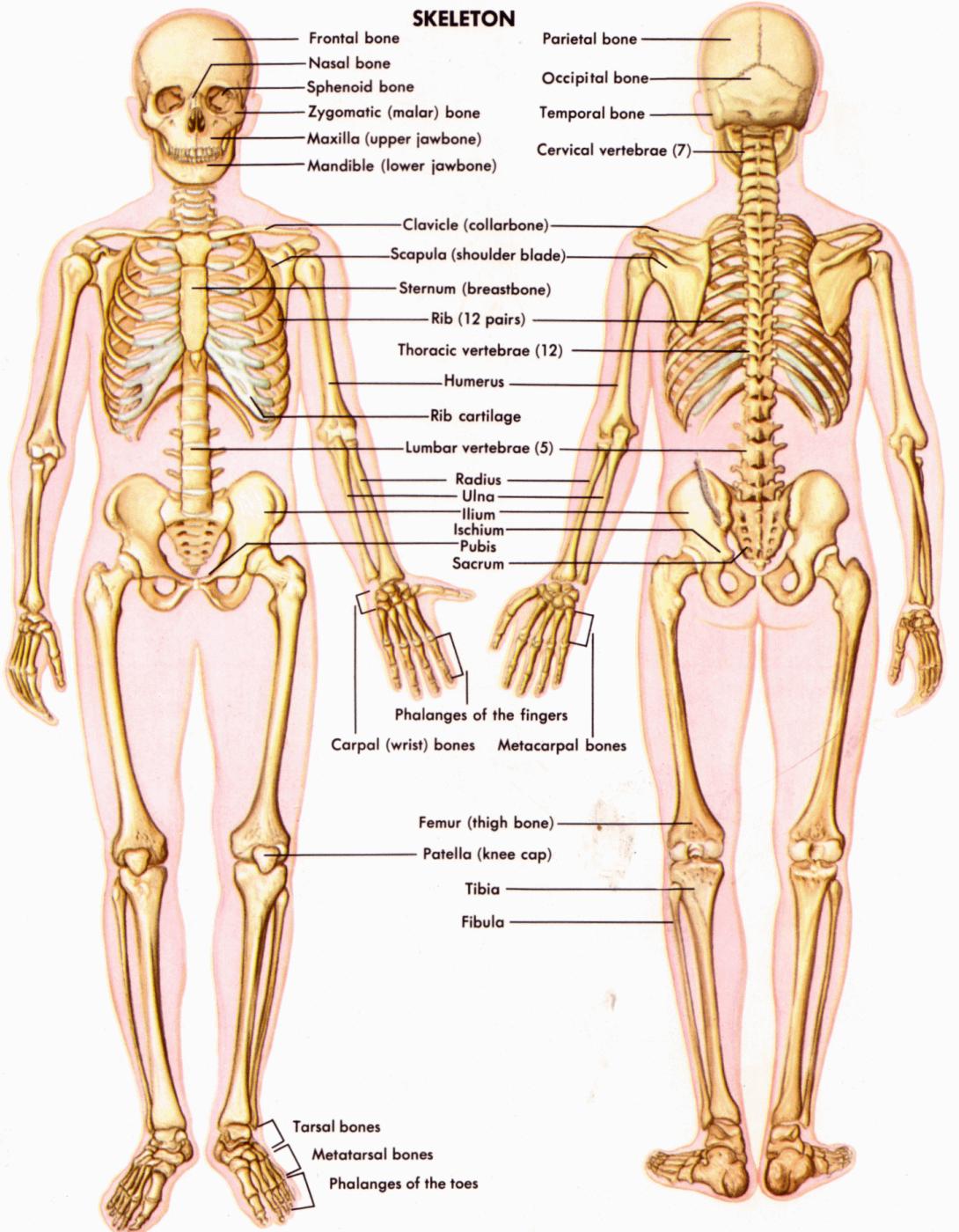
It takes 40-56 days for a cell to reach the epidermal surface

Grant thought:

Some drugs are administered by applying them to the skin. The drug diffuses through the epidermis to blood vessels in the dermis. What kind of substances can pass easily through diffusion? What kind have difficulty? Can drugs be *engineered* to pass through more easily??

*On your own: Look online for skin grafts

Skeletal System



Provides protection and support, allows body movements, produces blood cells, and stores minerals and fat. Consists of bones, associated cartilages, ligaments and joints

206 total bones

Skeletal System first to be “engineered”

Traditional route for skeletal system engineering: Bio-inert materials

Purposes:

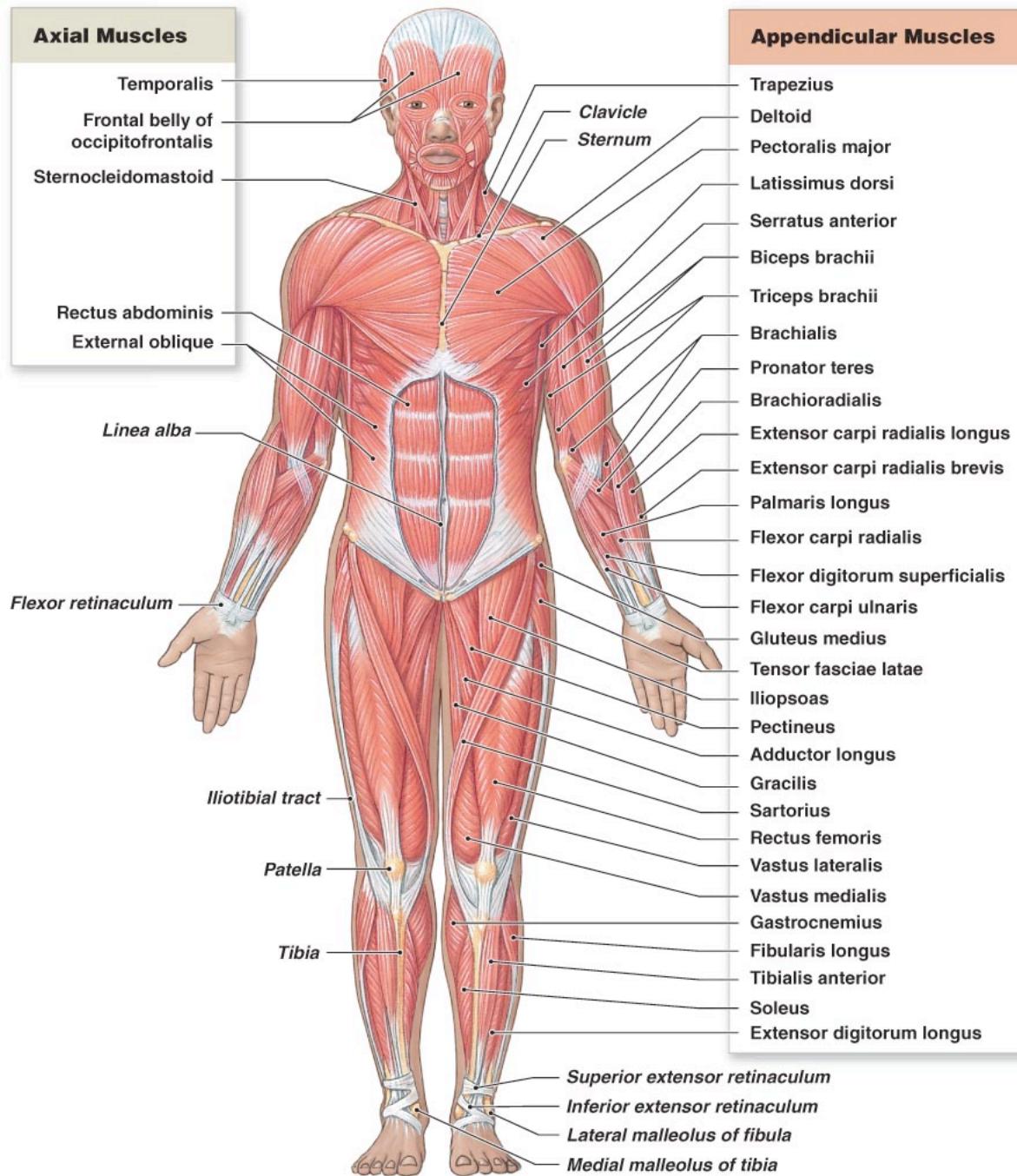
1. Replace tissue *structure and function*, not *bio-activity*
2. do not entice an immune response once implanted into the body.
3. Have incredible mechanical toughness withstand physiological loading
4. Long lasting in the body (won't degrade over time)

Metals	Ceramics	Polymers
316L stainless steel	Alumina	Ultra high molecular weight polyethylene
Co-Cr Alloys	Zirconia	Polyurethane
Titanium	Carbon	
Ti6Al4V	Hydroxyapatite	

Applications:

Skeletal tissue prosthesis (hip, knee replacement)



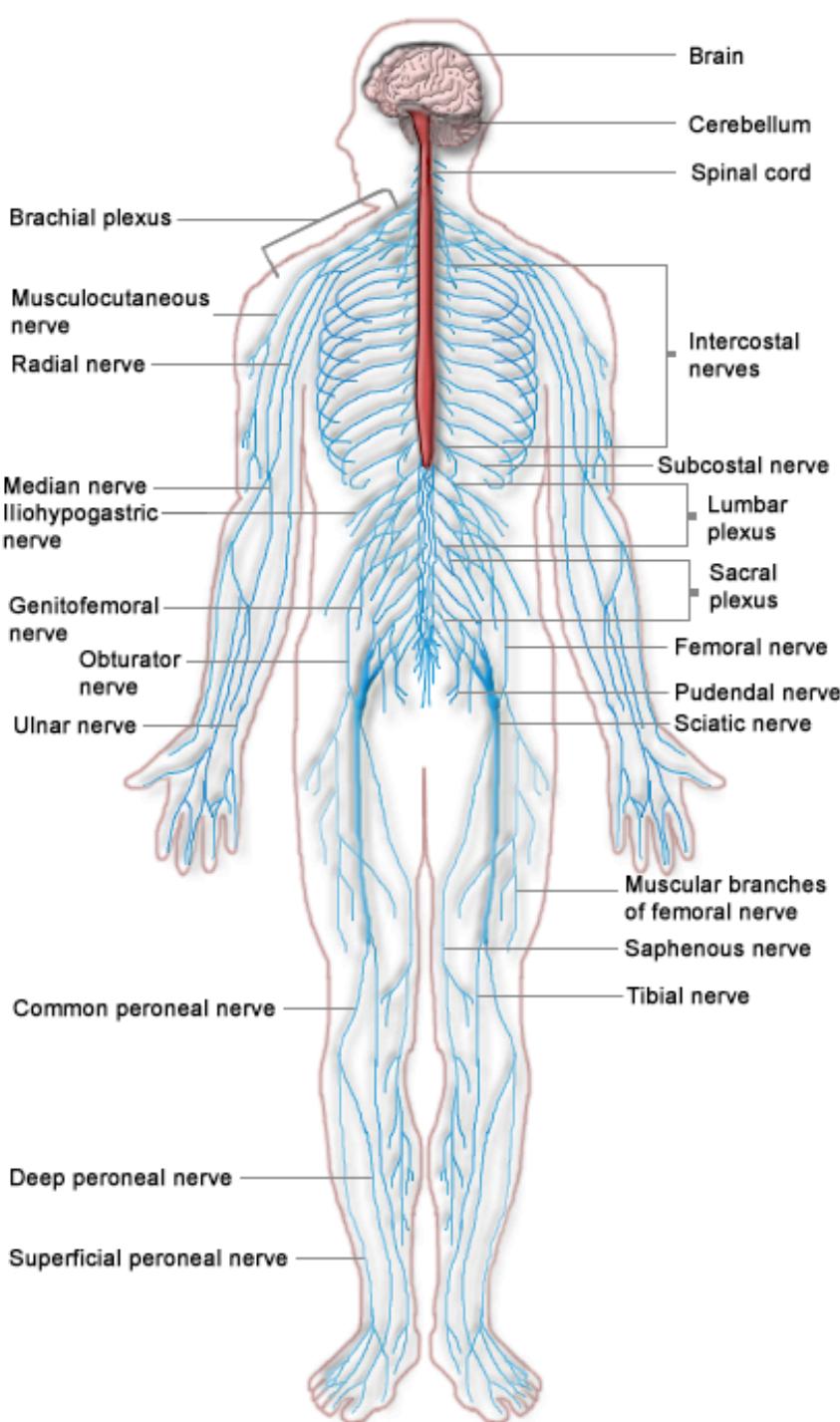


Muscular System

Produces body movements, maintains posture, and produces body heat. Consists of muscles attached to the skeleton by tendons

GRANT THOUGHT

Not truly engineered yet.
Many working on stem cell-to-muscle cell differentiation techniques.

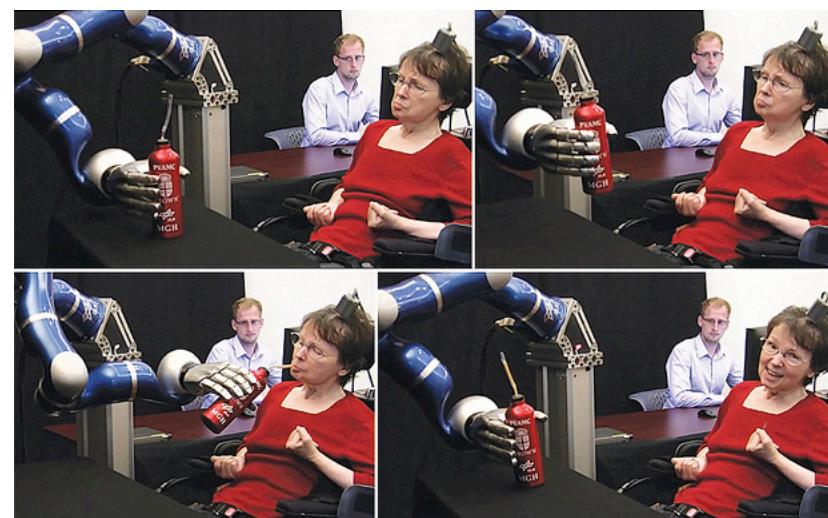


Nervous System

A major regulatory system that detects sensations and controls movements, physiologic processes, and intellectual functions. Consists of the brain, spinal cord, nerves, and sensory receptors.

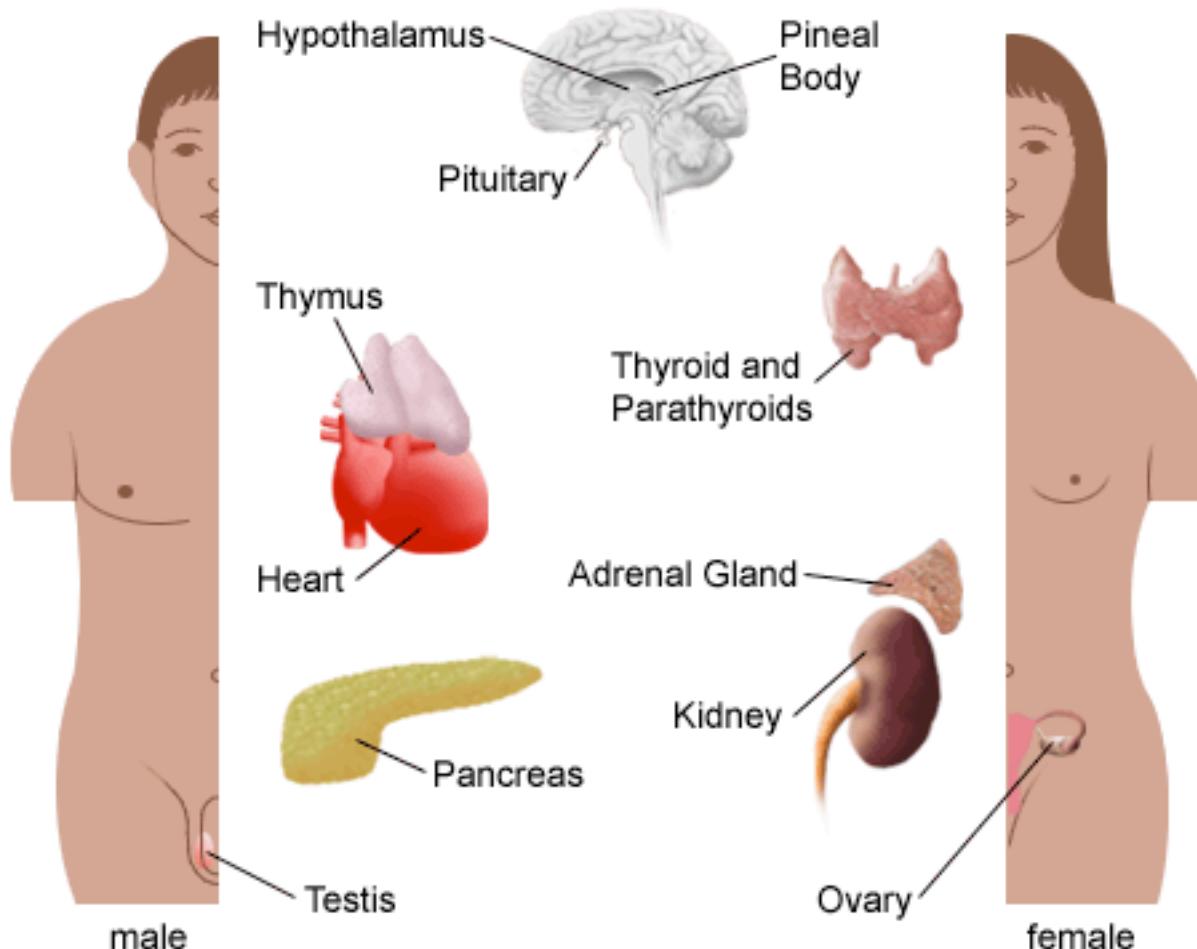
Engineering advances:

1. Functional electrical stimulation: use outside or implanted devices to stimulate otherwise inactive nerves.



Endocrine System

Endocrine System

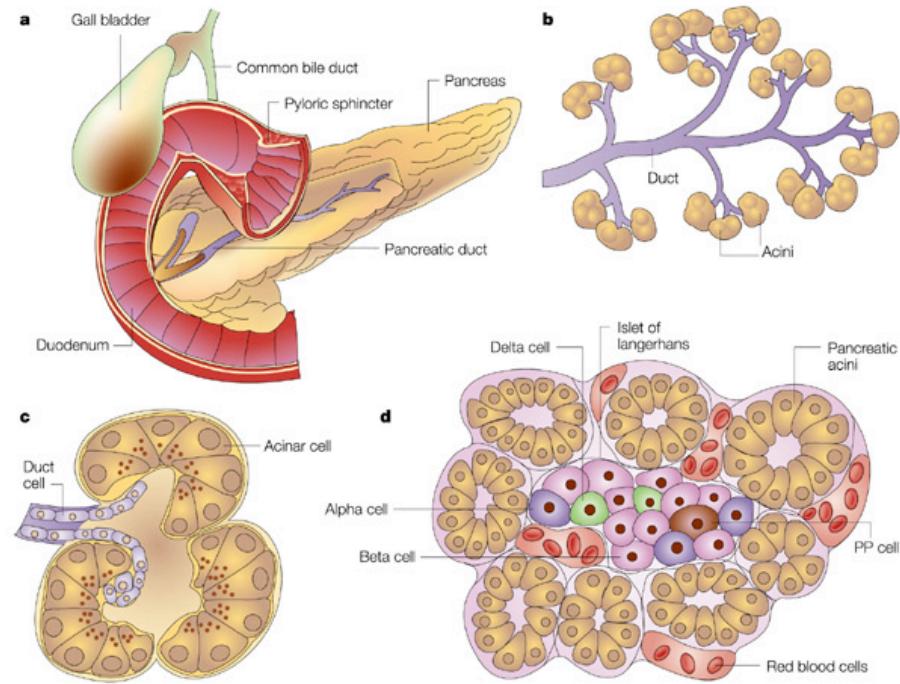
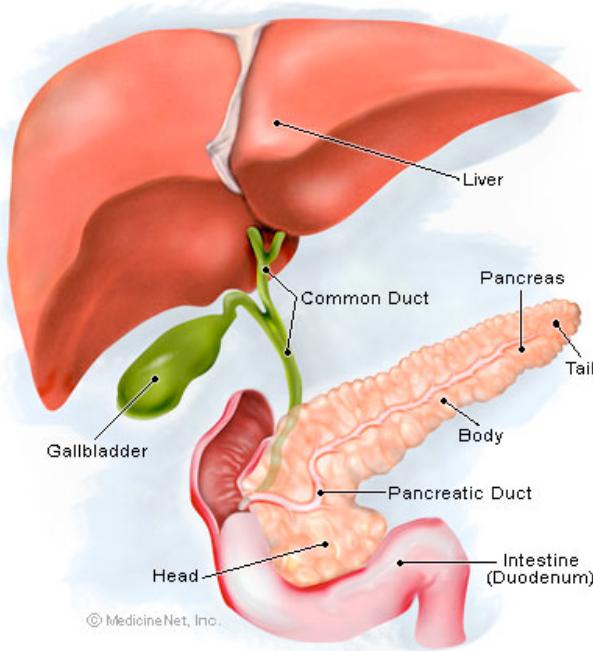


GRANT THOUGHT

Not truly engineered yet.
Typically treated with repeated injections of hormones or daily pills.
Can be adapted to a patch or implanted, permanent device?

A major regulatory system that influences metabolism, growth, reproduction, and many other functions. Consists of glands, such as the pituitary, that secrete hormones

Pancreas Function

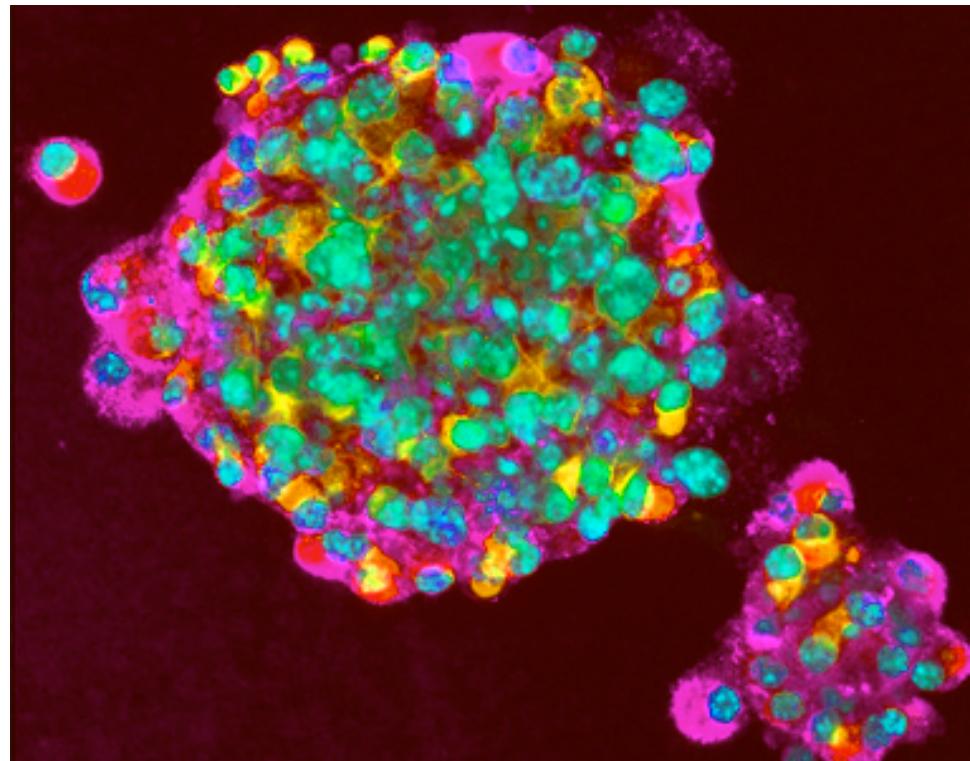


Produces many enzymes (including insulin) to break down food (carbohydrates)
Secretes enzymes into the small intestine after measuring carbohydrate levels.
Different islet cells produce different enzymes.

Hydrogel culture of islet cells

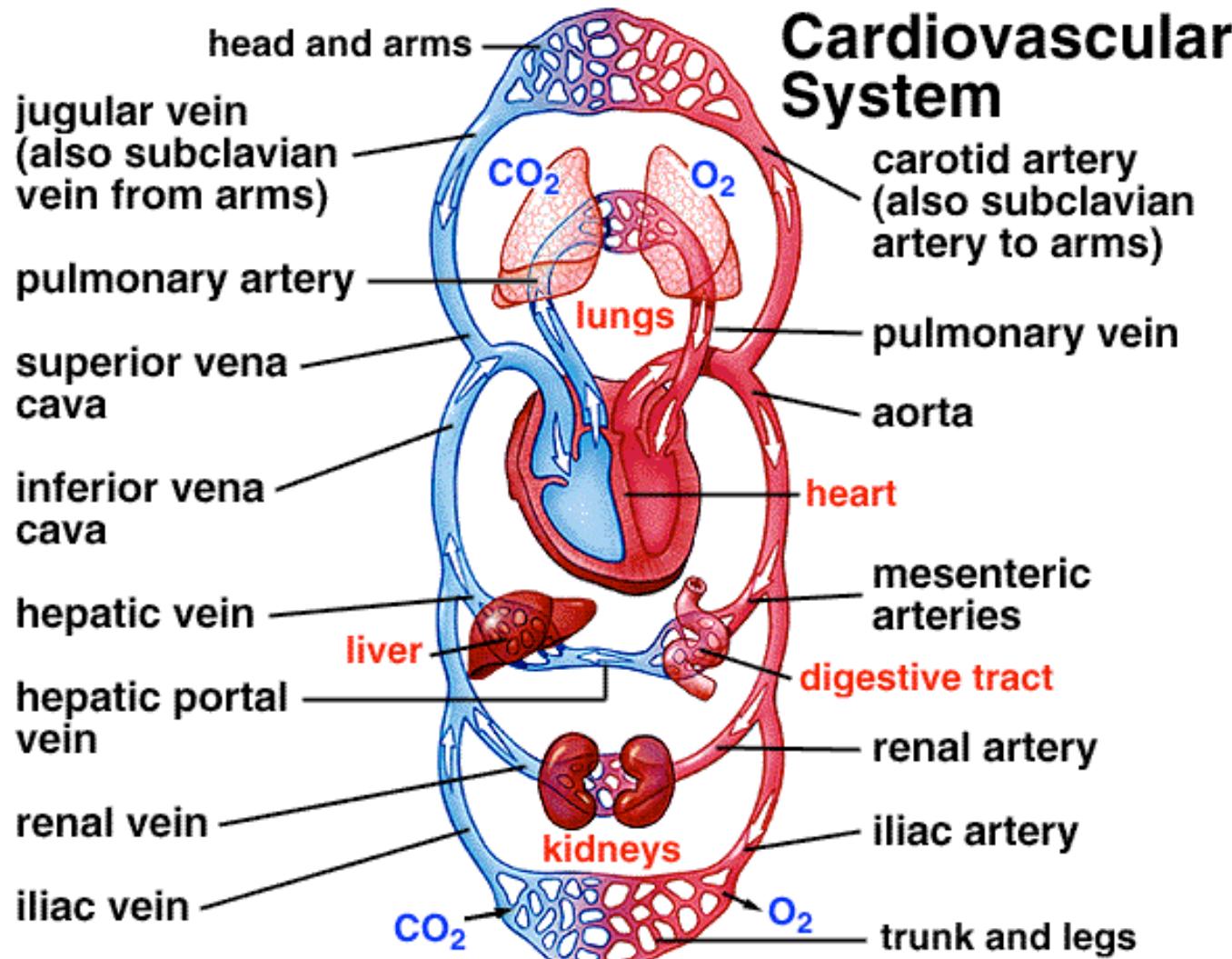


Stem cell differentiation to islet cells



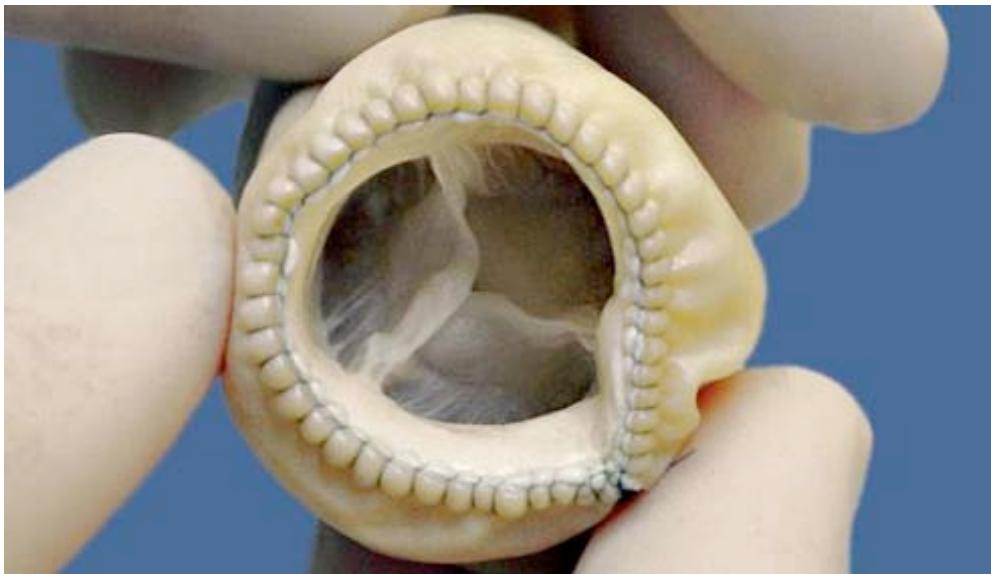
Cardiovascular System

Sylvia S. Mader, Inquiry into Life, 8th edition. Copyright © 1997 The McGraw-Hill Companies, Inc. All rights reserved.

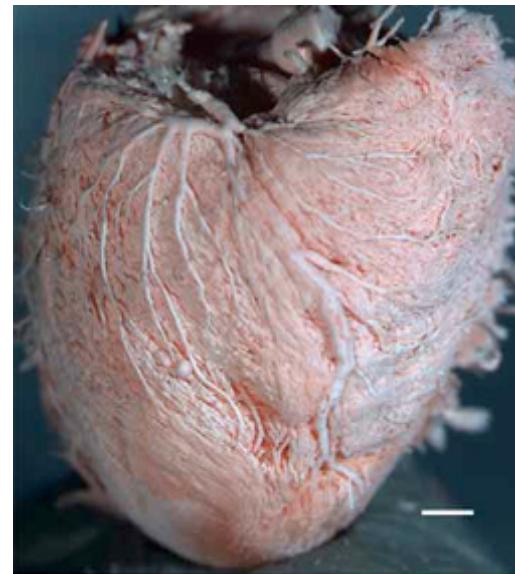


Transports nutrients, waste products, gases, and hormones throughout the body; plays a role in the immune response and the regulation of body temperature. Consists of the heart, blood vessels, and blood.

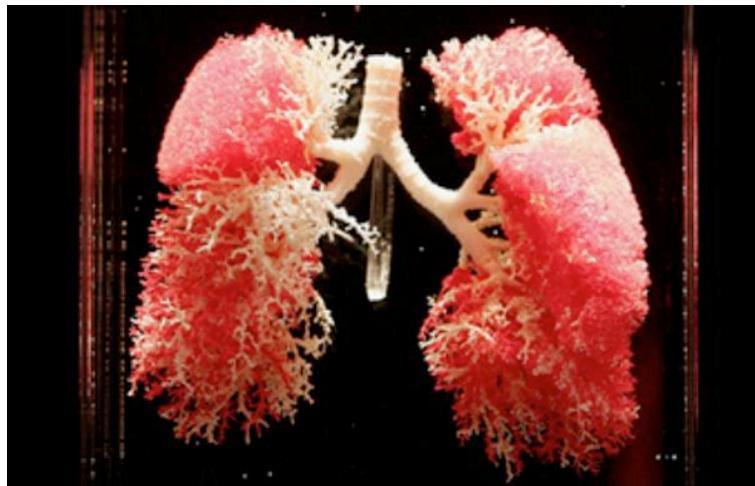
Lots of active TE in cardiovascular area!



Heart Valves, from cadavers or animals



Rat heart



lungs

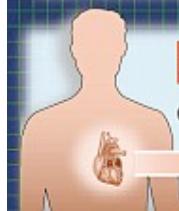
HOW TO MAKE A HEART



1 Heart removed from donor. Washed in detergent to remove heart cells



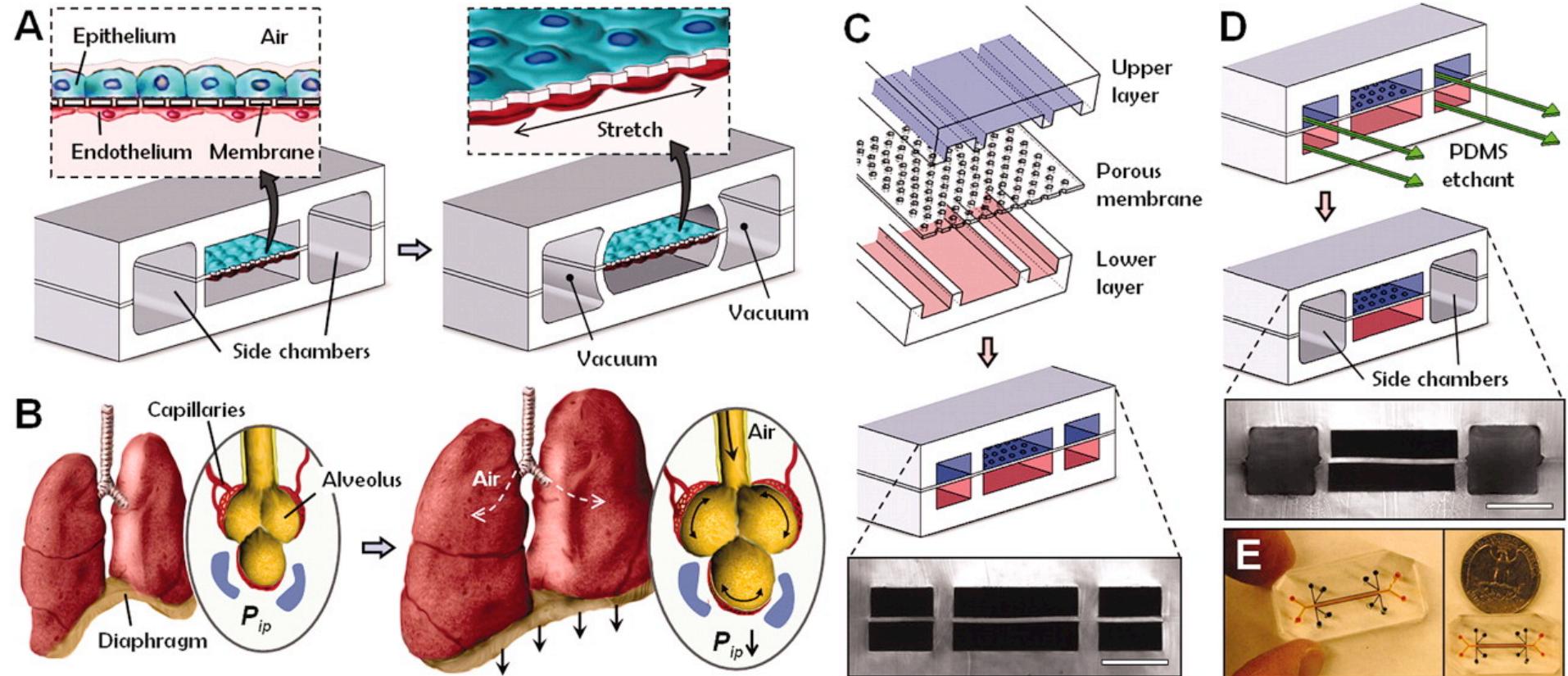
2 Collagen 'skeleton' of heart left behind



3 Stem cells taken from patient and grown in a dish of nutrients

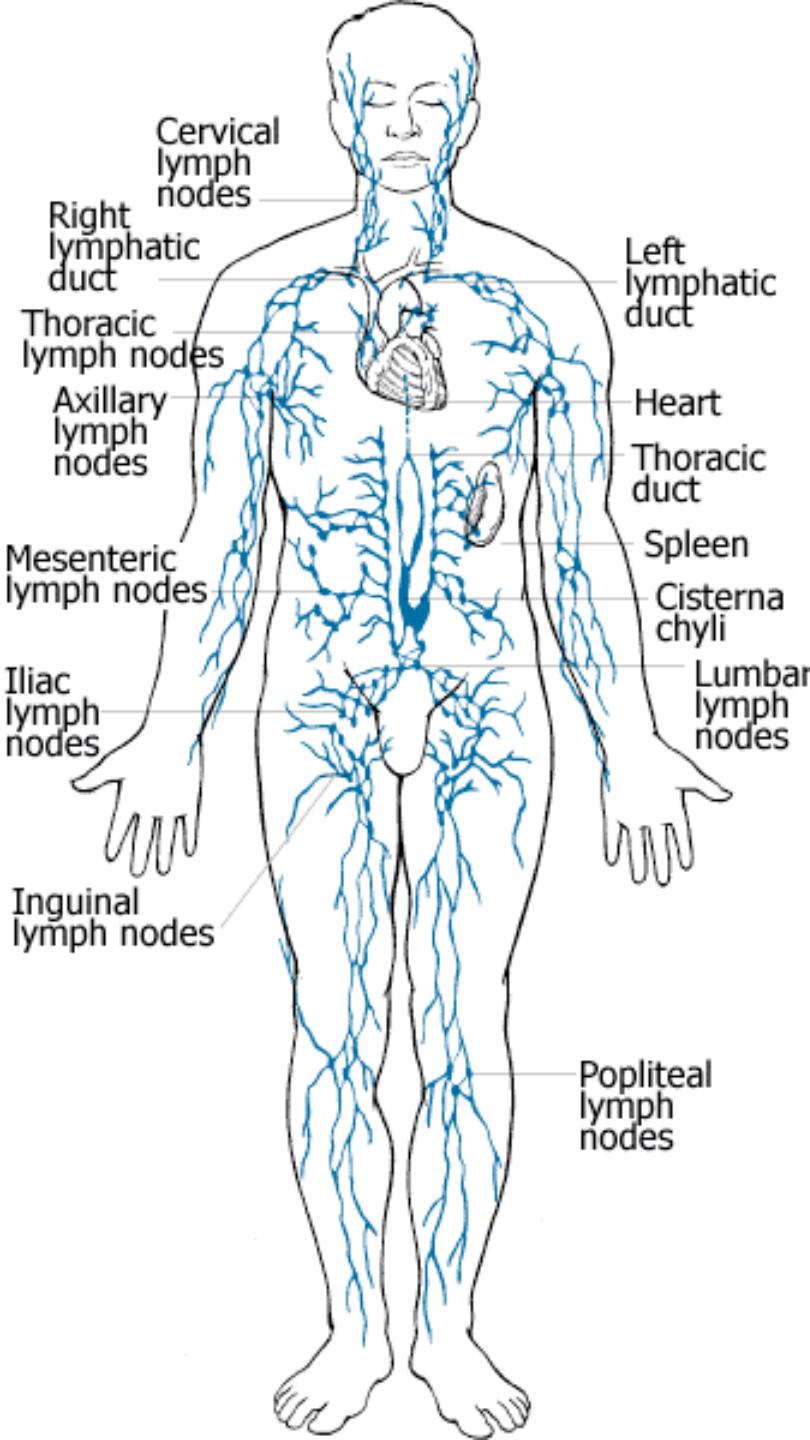


Preview to wiki? Organs on a chip



Lung on a chip

Weiss Institute, Harvard

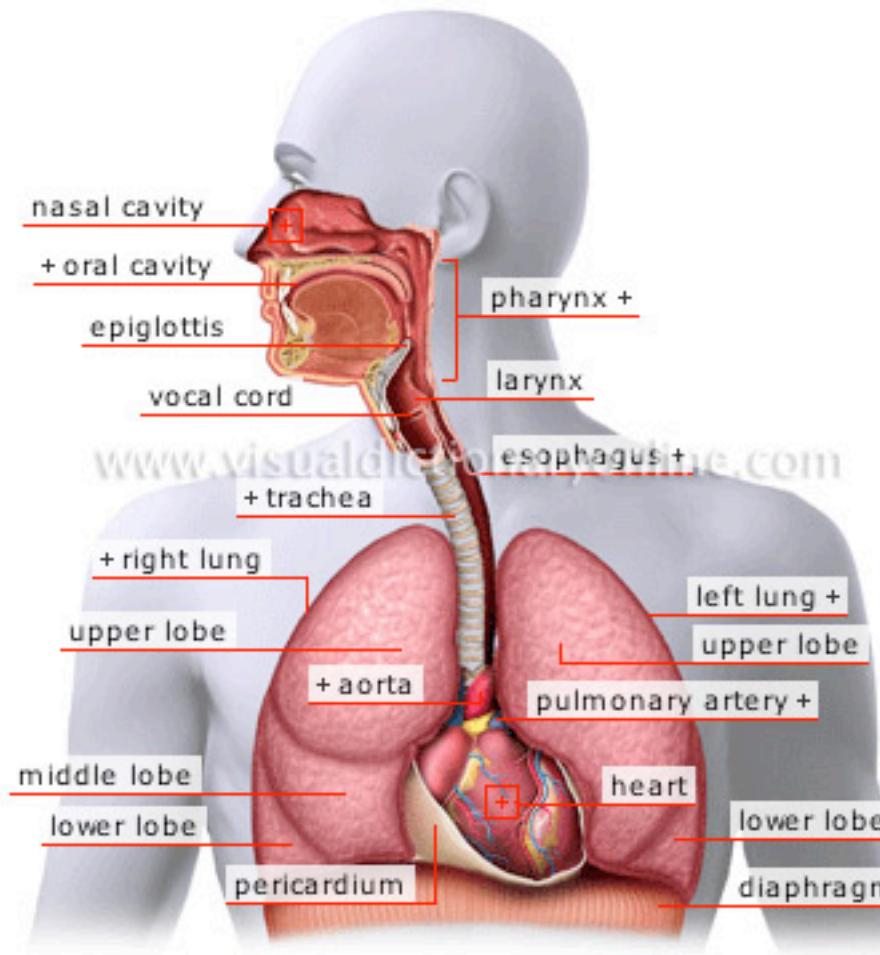


Lymphatic System

Removes foreign substances from the blood and lymph, combats disease, maintains tissue fluid balance, and absorbs fats from the digestive tract. Consists of the lymphatic vessels, lymph nodes, and other lymphatic organs.

Only bioengineer (Chemical Engineer) working on this is Melody Swartz, EPFL. She studies flow patterns to the lymph using principles of fluid dynamics.

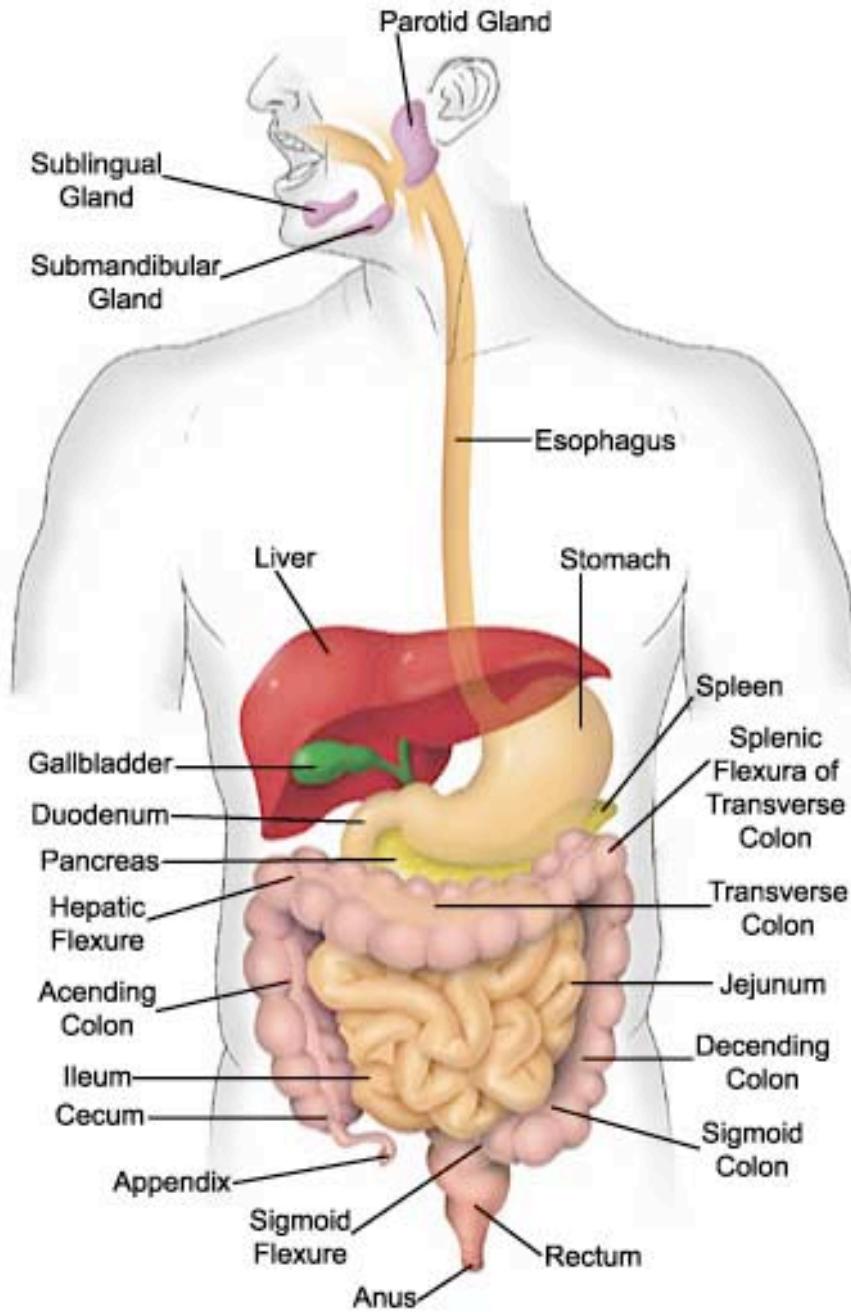
Respiratory System



Same engineering principles as cardiovascular system

Exchanges oxygen and carbon dioxide between the blood and air and regulates blood pH. Consists of the lungs and respiratory passages.

Digestive System



Performs the mechanical and chemical processes of digestion, absorption of nutrients, and elimination of wastes. Consists of the mouth, esophagus, stomach, intestines, and accessory organs.

Teeth are part of the digestive system – lots of obvious TE activity w.r.t dental implants.

Some work in liver. Check out Linda Griffith and Sangeeta Bhatia (both at MIT).

Artificial Liver Device

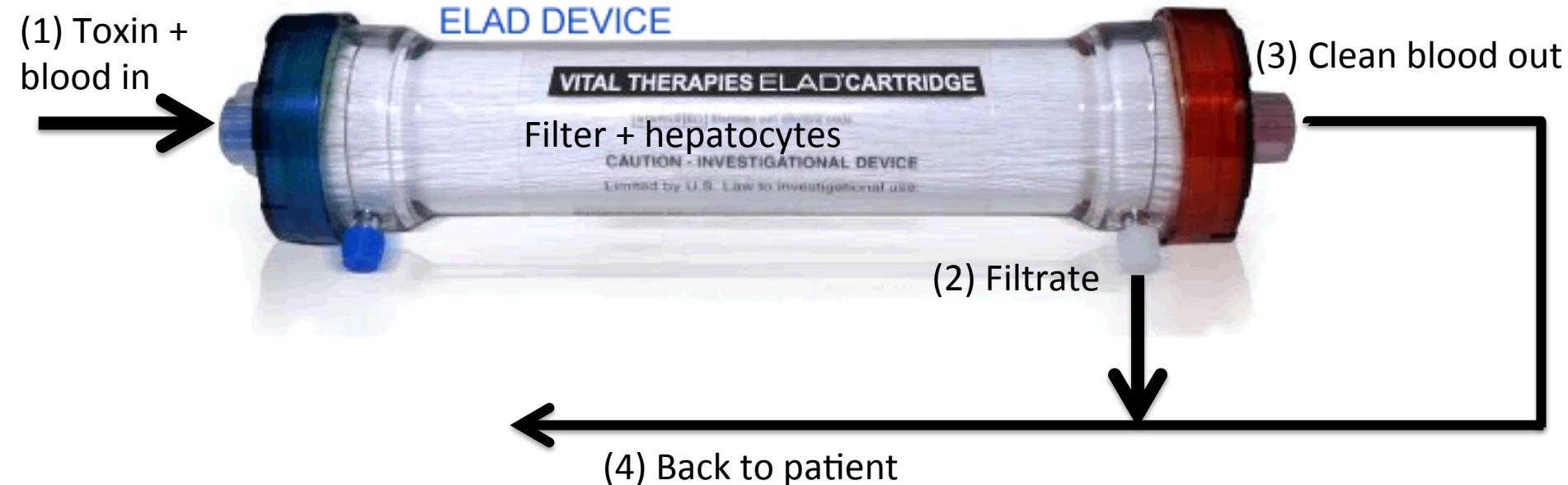


Healthy

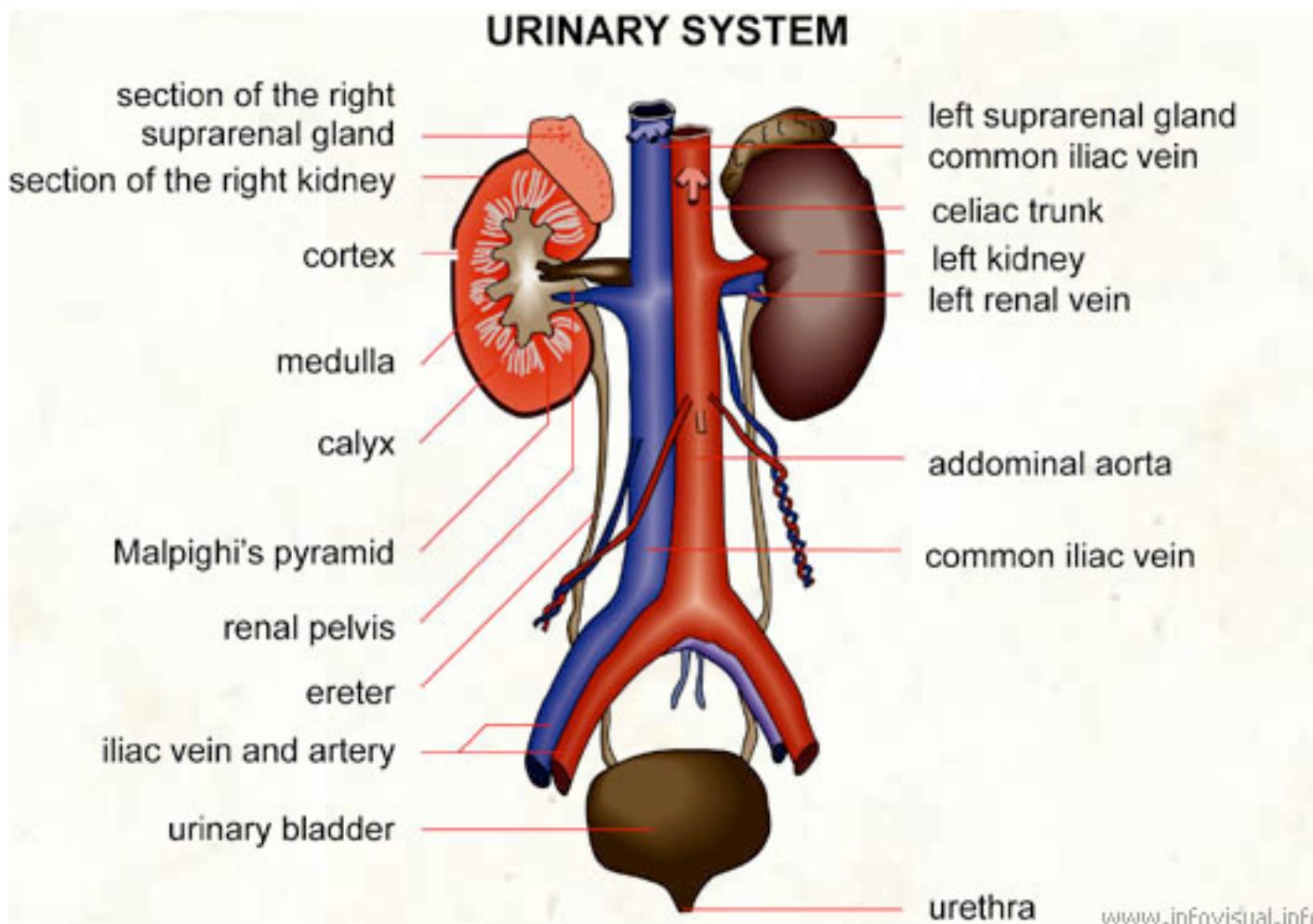


Cirrhosis

Synthetic Filter: retains very large compounds (>100,000 g/mol)
+ Hepatocytes: process bilirubin (reaction)



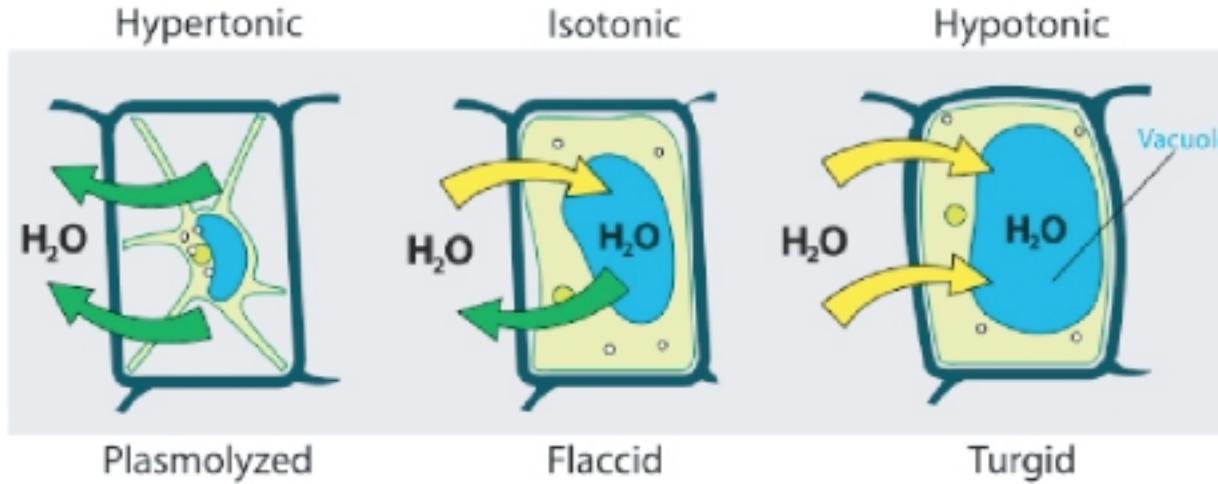
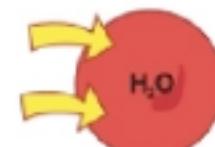
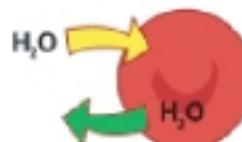
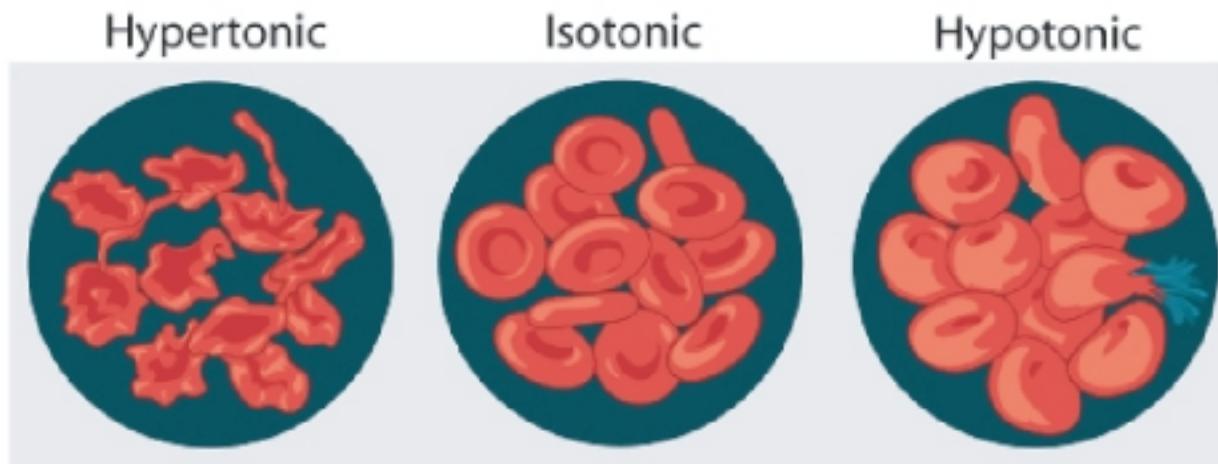
Urinary System



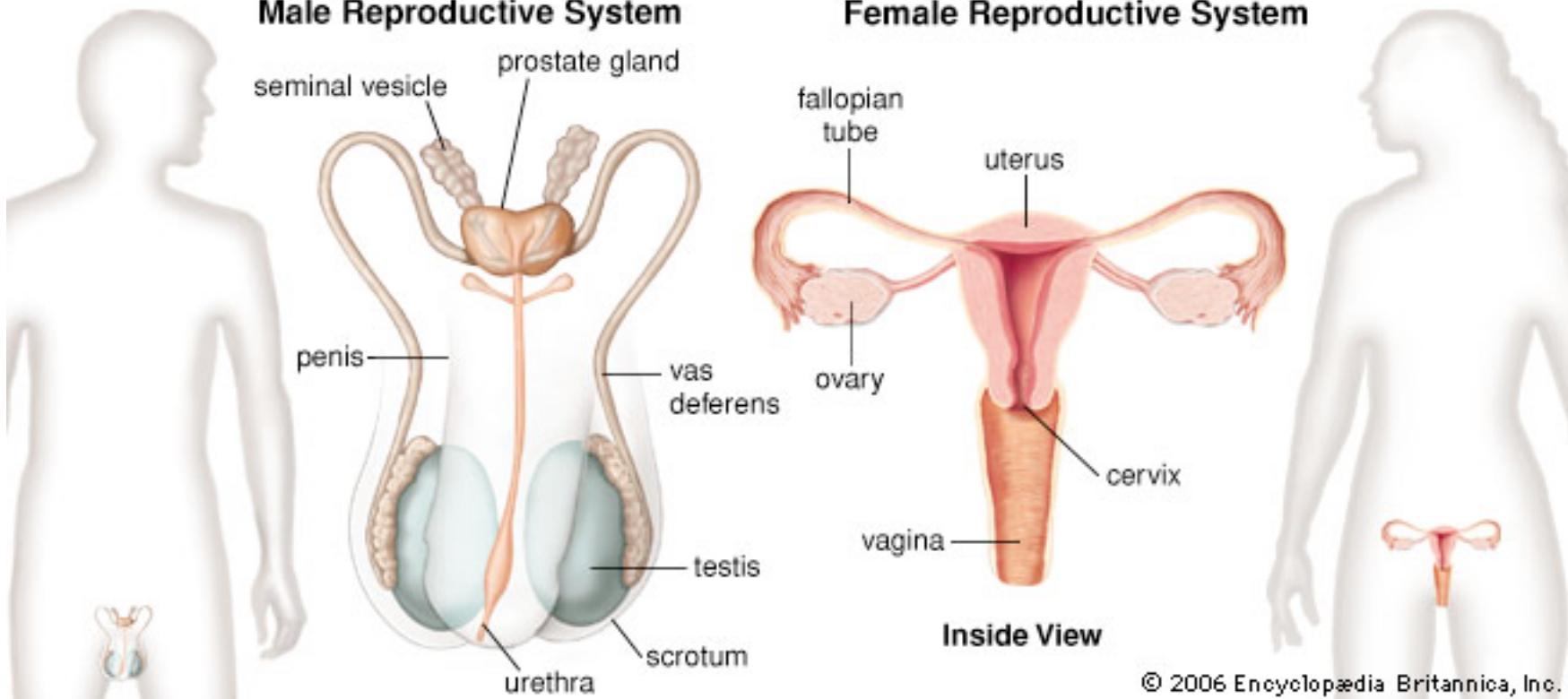
Removes waste products from the blood and regulates blood pH, ion balance, and water balance. Consists of the kidneys, urinary bladder, and ducts that carry urine.

Tony Atala (wiki option) – has built replacement bladders.

Why is the salt balance important?



Reproductive System



F: Produces oocytes and is the site of fertilization and fetal development; produces milk

M: Produces and transfers sperm cells

Both: produces hormones that influence sexual functions and behaviors.

Tony Atala: lots of tissue engineering of reproductive system organs.

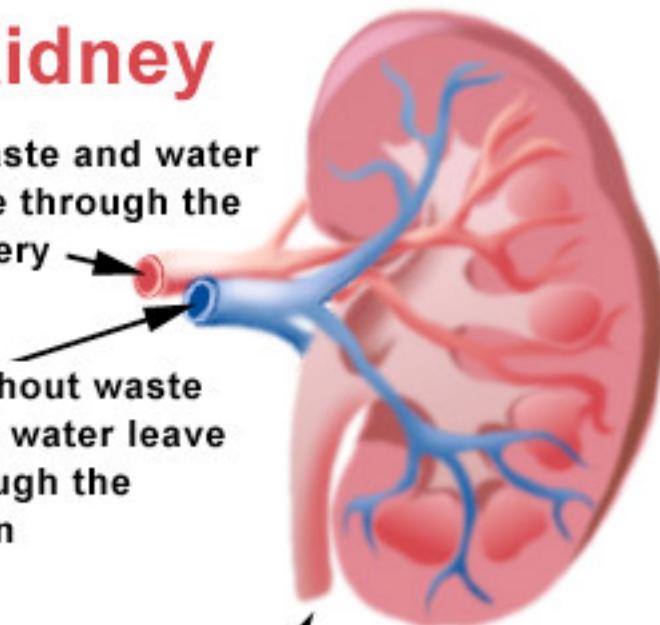
Kidney Function

A Kidney

Blood, waste and water enter here through the Renal Artery

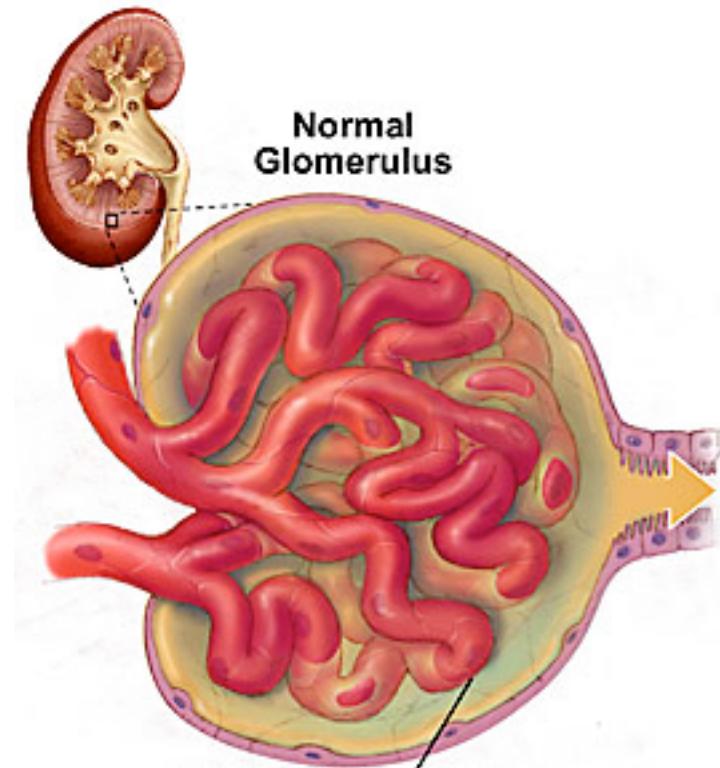
Blood without waste or excess water leave here through the Renal Vein

Excess water and Toxic Waste in the form of Urine leaves here via the Ureter



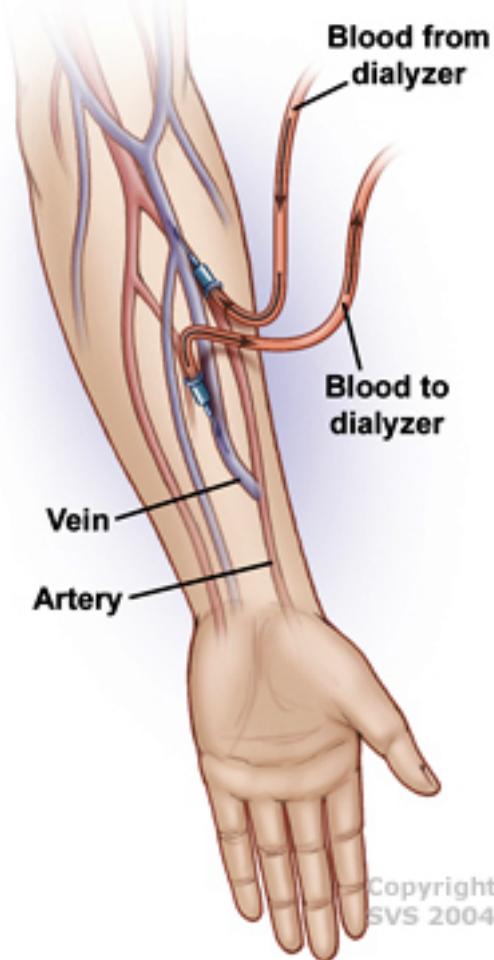
Normal Glomerulus

Blood with impurities is filtered creating urine

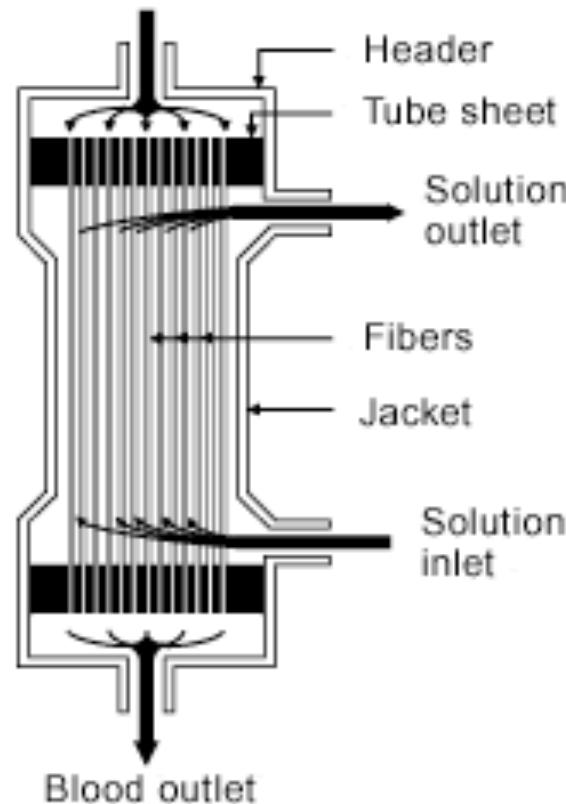


Dialysis given to patients with kidney failure

Arteriovenous fistula

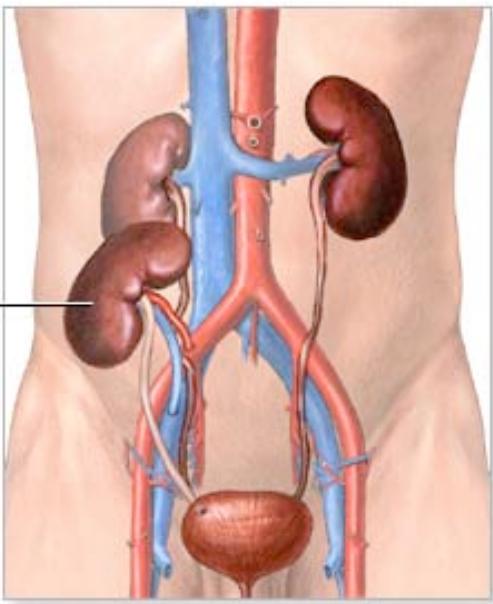


Blood inlet



Permanent solutions to kidney failure

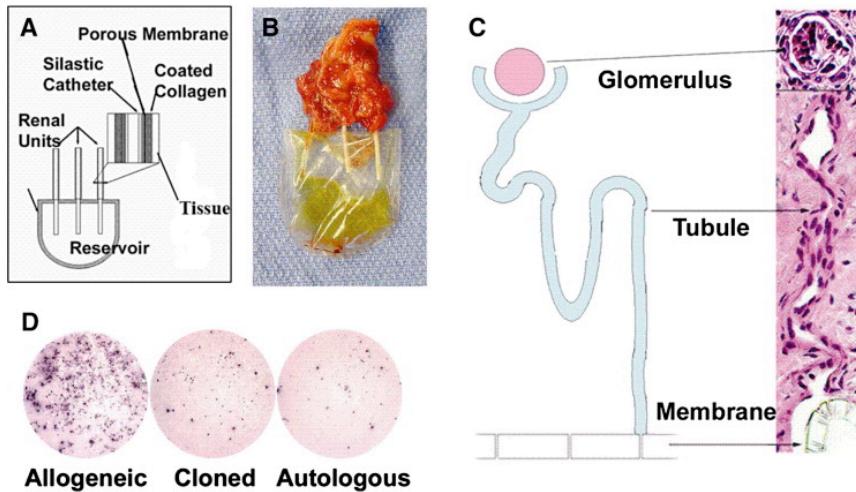
Transplanted kidney



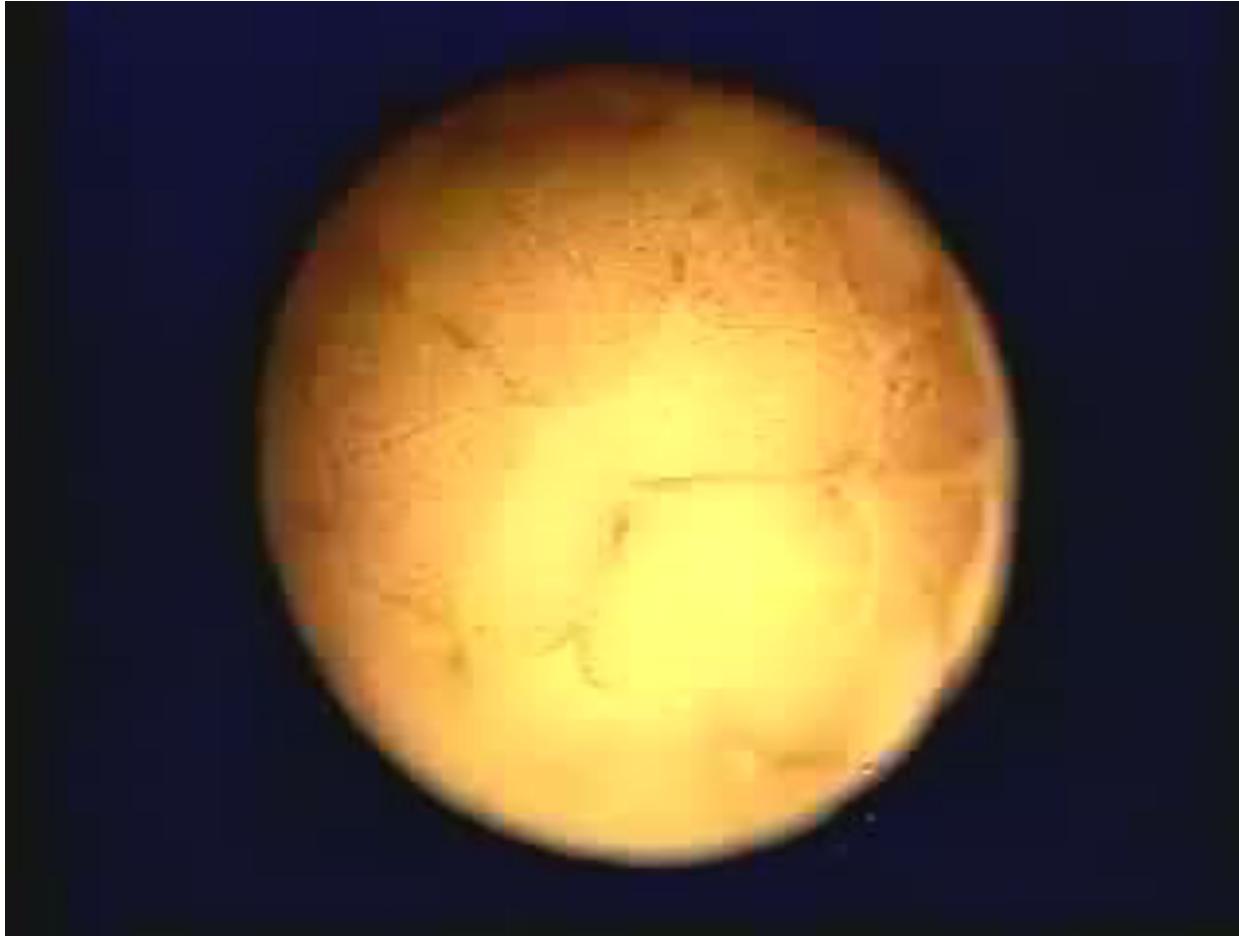
Decellularized kidney



Tissue engineering bio-hybrids?

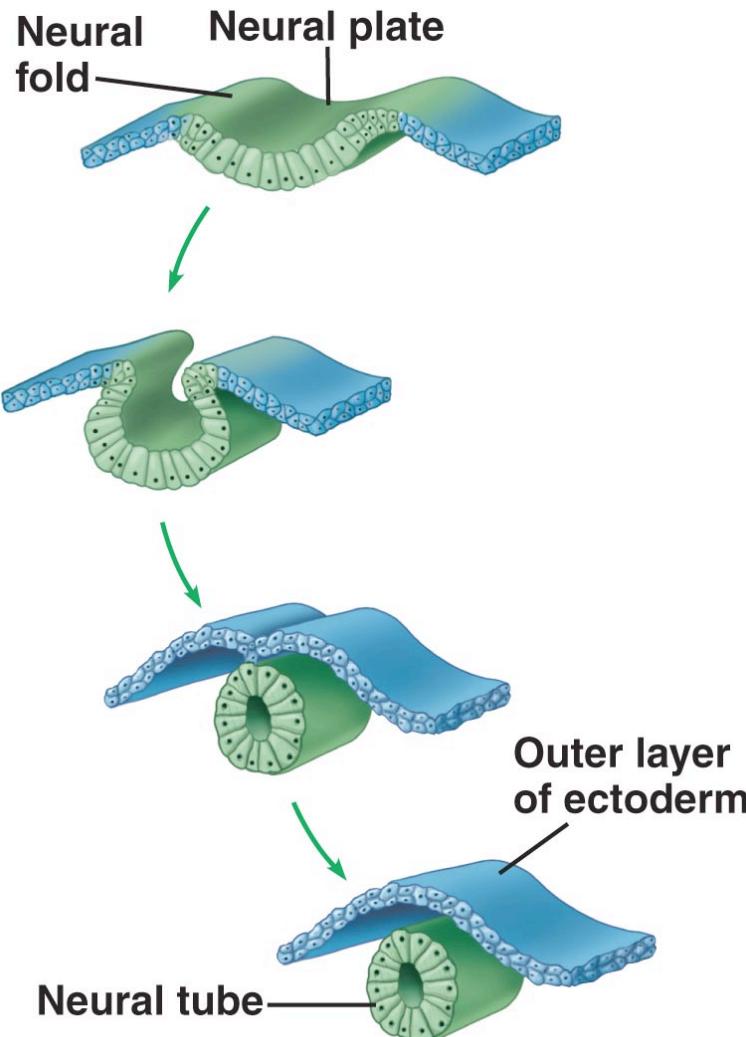


How do all these organ systems develop?



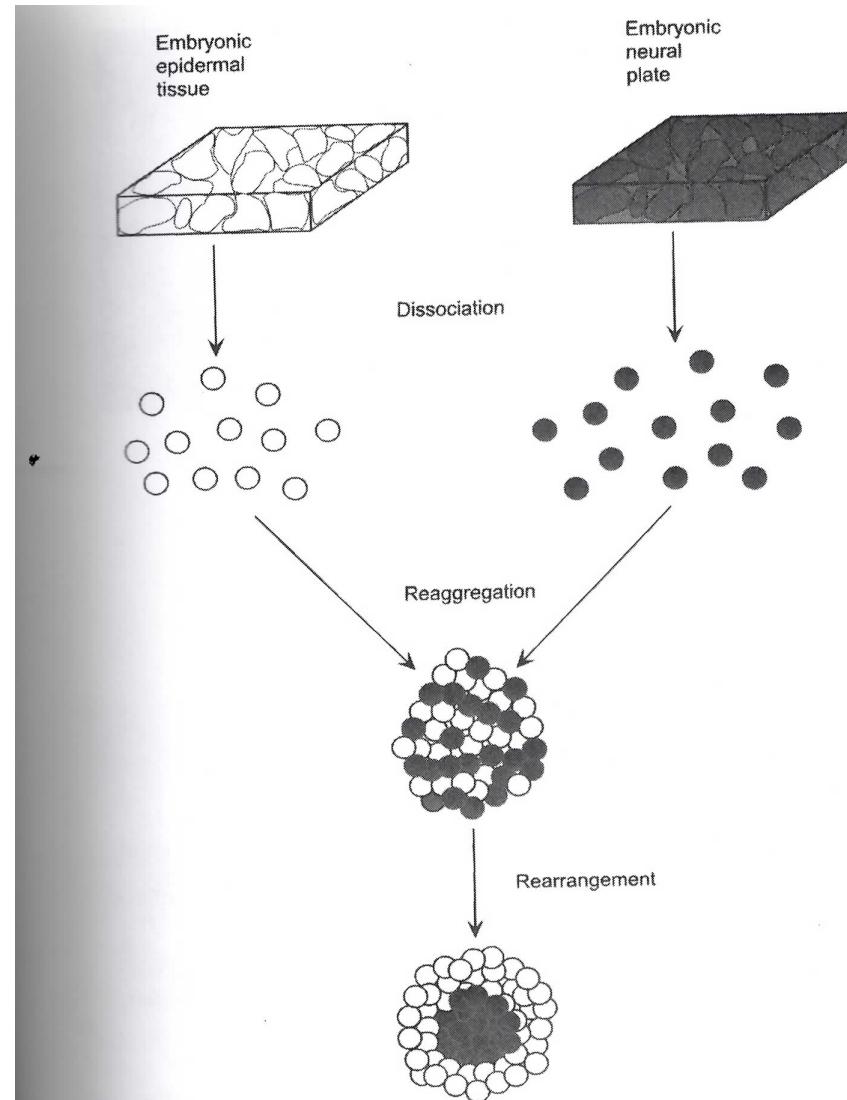
MBOC: xenopous egg

Self-sorting in embryonic cells captures some of developmental process

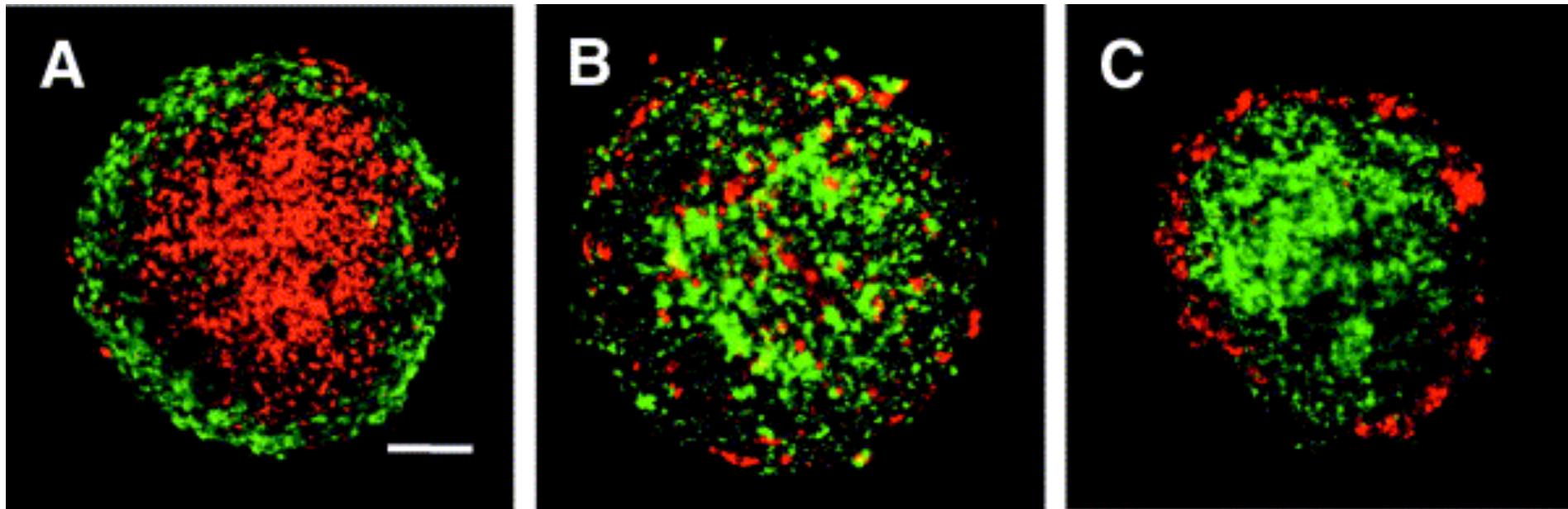


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Neural plate formation in frog embryo development



Chemical Equilibria of Cell Sorting



From Duguay et. al., *Developmental Biology*

This type of approach can sometimes get an engineered tissue “started” – to capture tissue function naturally.

Grant thought: manipulate cell surface receptors to get them to self sort??