

## 3. PŮVOD TKÁNÍ A JEJICH ROZF

Tkáň lze definovat jako soubor morfolos se shodnou nebo velmi podobnou funkční mi složkami orgánů lidského těla.

Tkáně se vyvíjejí ze zárodečnýc togeneze v průběhu embryonál entoderm a mezoderm a Mezenchym je embry ektodermu) a vyplň síťovitou textur Tkáně se tkáň epit

Tk/

sebou je Epitelov. střevo), sekrec. i kontrakcí (např. 11. Vzhled a stavba kon

#### Tkáň pojivová

Pojivová tkáň je mezenchymo Skládá se z buněk a mezibuněčné hn. ... a amorfní hmotu základni.

# Část II. Čtyři základní typy u.

#### Epitelová tkáň

#### CÍLE STUDIA

- Tato kapitola by měla studentoví pomocí poznat čtyři základní typy tkání
- poznat ciyri zakiadni typy tkani poznat strukturální a funkční charakteristiky, které odlišují epitelovou tkáň od dalších tří záklac
- men typu tкаnn poznat typy epitelové tkáně a uvést příklady míst, kde se jednotlivé typy mohou nacházet poznat typy epiteiove ikane a uvesi pinkiauy inisi, kue se jeunomve typy monou nacinazer poznat funkční vlastnosti každého typu epitelové ikáně a uvést jejich vztah ke struktuře tkáně od se poznat su se poznat nacinazer родна нивели унваноми какиено тури ерненоve ткане a uvest jejich vztan ке мижине ткане popsat speciální funkce jednotlivých typů epitelových buněk a uvést příklady míst, kde se jednot-
- na mikrofotografiich poznat epitely a určit jejich funkci podle struktury a lokalizace znát kritéria, která se užívají při klasifikaci žláz
- znát druhy žláz u člověka a uvést příklady míst, kde se mohou nacházet na mikrofotografiích a schématech poznat zlázy a určit jejich typ

## MAX-Yield<sup>TM</sup> OTÁZKY KE STUDIU

- Vyjmenujte hlavní funkce epitelových tkání (II.A.¹).
- Ze kterého(ých) embryonálního(ích) zárodečného(ých) listu(ů) se epitelové tkáně vyvíjejí? Uveďte Ze kierenojych emoryonammojrch zarouechenojych usutu) se epherove ikane vy příklady epitelů odvozených od jednotlivých zárodečných listů (II.H.; tabulka 4-1).
- prakacy epitetu odvozenych od jednoutvych zaroucchych ustu (H.Fr., taounka +-1). Vyjmenujte strukturální a funkční charakteristiky epitelových tkání, které je odlišují od ostatních ypinenujte strukturami a iunkeni enarakteristiky epitejových tkani, které je odiisují od ostatních vpů tkání. Vezměte v úvahu polaritu buněk (IV.), specializace apikálních (IV.A.), laterálních ури калі. vezmete v uvanu poiaritu ounek (Iv.), specianizace аріканісці (Iv.т.), насізанісці (IV.B.) a bazálních (IV.C.) povrchů, způsob výživy (II.F.) a intenzitu mitotického dělení (II.E.). Popište bazální laminu s ohledem na její lokalizací, složení a barvící vlastnosti (IV.C.)

Které struktury a molekuly pomáhají připevnit enitelová knálvy

Histologie (z řeckého histos = tkáň, logia = studium) je nauka o stavbě tkání. Tkáně lze chymu (derivát mesodermu). Vyznačuje se hojnou definovat jako komplex morfologicky podobných buněk, specialisovaných k výkonu určité funkce. jsou uloženy rozličné typy buněk, plnící řadu Jsou materiálem pro stavbu orgánů těl mnohobuněčných organismů, metazoí. Za embryonálního vývoje jedince (ontogenese) se tkáně díferencují ze 3 mového. Tvoří ji buňky nebo syncytium. Její elezárodečných listů, ektodermu, entodermu a mesodermu, procesem zvaným histogenese. Na jejím podkladě vznikají čtyři základní typy tkání: i pohyb orientovaný v příslušném směru.

 Tkáň epitelová – vzniká ze všech tří zárodečných listů. Tvoří ji buňky těsně k sobě přiložené s malým množstvím mezibuněčné hmoty. Uspořá-F. Hety legifol poyrchy nebo v enitelové

2. Tkáň pojivová, podpůrná – pochází z mesenúčastí mezibuněčné základní hmoty, ve které

3. Tkáň svalová – je původu převážně mesodermenty jsou protáhlého tvaru. Jejich cytoplasma je opatřena prvky, které umožňují její kontrakci, a tím

4. Tkáň nervová – pochází z ektodermu. Její nejvýznamnější komponentou jsou nervové buňky neurony, schopné vytvářet nervový vzruch a předávat jej z buňky na buňku.

#### ely)

a budou proto probrány zde, ackoliv lze prokázat v různé míře a zastouostatních typů tkání.

#### 1ezibuněčné spoje, kty epitelových buněk

pitelových buněk je podmíněna speciaemy sousedních buněk ve struktury, bezpečena jejich kohese.

volného povrchu buněk je interceluutěsněna tzv. tmelovými lištami. Lze impregnací roztokem solí stříbra, poením železitým hematoxylinem podle či jinými metodami. Na řezu vedežně s povrchem buněk vytvářejí tmeaz šestiúhelníku. Na řezech kolmých tmelové lišty patrny jako tmavé body n povrchu buněk (obr. 67).

ovém mikroskopu byla tato specialisora popsána jako tzv. spojovací komplex, i složkami (obr. 64). Těsně pod povr-

κ v těle je trvale usedlá (fixní) a uspořádaná do souborů. Soubor stejně ných buněk spojených mezibuněčnými kontakty a mezibuněčnou hmotou nych ounex spojenych meziounechynh kontakty a meziounechou mitotou (káň. Rozlišujeme čtyři základní typy tkání: epithely, pojiva, svalovinu a tkáň

pithely. Jsou to soubory buněk s četnými vzájemnými kontakty a minimem mezibuněčného prostoru. Základní dělení: krycí epithely, žlázové epithely. Pojivové tkáně. Stavební princip: málo buněk, větší mezibuněčný prostor vypl-

něný mezibuněčnou hmotou (např. kolagenní a elastická vlákna, proteoglykany, neny meziourectora minorou mapr. Adagemin a ciasticka viakita, proteogrykany, minerály). Její uspořádání je rozhodující pro specifické biomechanické vlastnosti jednotlivých typů pojivové tkáně. Základní dělení: řídké a tuhé kolagenní vazivo, slachy, ligamenta, tukové vazívo, chrupavka, kost.

ávod

Nervová tkáň. Soubor nervových buněk včetně jejich výběžků a gliových buněk; je specializována na přenos a zpracování informací, které jsou založeny na elektro-

Tkáň svalová. Je to soubor buněk schopných koordinovaných, makroskopicky patrných kontrakcí. Rozčlenění: příčně pruhované svalstvo (kosterní a srdeční), hladká svalovina.

**Orgán** je vždy tvořen z většího počtu tkání. Tkáň specifická pro orgán – většinou epithel – se označuje jako parenchym, na rozdíl od vazivového stromatu, které poskytuje organům mechanickou soudržnost a ve kterém jsou uloženy cévy (krevní a lymfatické) a nervy. Původ různých typů tkání a orgánů ze tří zárodečných listů (ektoderm, mesoderm, entoderm) mladého embrya je rekapitulován na str. 447.

#### OV

anickou,

ioxám, aj.).

ruhovaná koster.

ζáň

#### Tissues are aggregates or groups of cells organized to perform one or more specific functions.

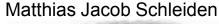
At the light microscope level, the cells and extracellular components of the various organs of the body exhibit a recognizable and often distinctive pattern of organization. This organized arrangement reflects the cooperative effort of cells performing a particular function. Therefore, an organized aggregation of cells that function in a collective manner is called a tissue [Fr. tissu, woven; L. texo, to weave].

Although it is frequently said that the cell is the basic functional unit of the body, it is really the tissues, through the collaborative efforts of their individual calls, that are more will

variations in general appearance structural organization, and physiologic properties of the val ious body organs, the tissues that compose them are classified into four basic types.

- Epithelium (epithelial tissue) covers body surfaces, link body cavities, and forms glands.
- Connective tissue underlies or supports the other this basic tissues, both structurally and functionally.
- Muscle tissue is made up of contractile cells and responsible for movement.
- Nerve tissue receives, transmits, and integrates into mation from outside and inside the body to control to activities of the body.

#### MODERN CELL TEHORY



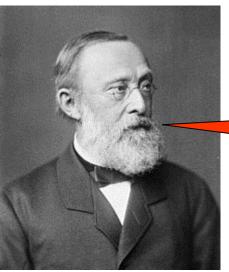


- Cells are the basic units of any organism
- New cells origin only from other cells
- •
- •Cells **exchange energy** (open thermodynamic system)
- Genetic information is inherited in new generations
- Chemical and structural composition of cells is generally identical



J.E.P.

**Rudolf Wirchow** 

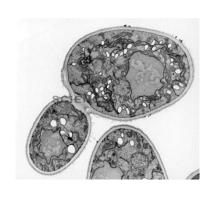


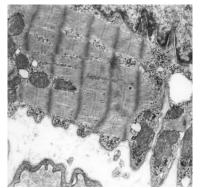
Omnis cellula e cellula!

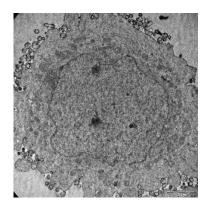




#### CELL AND TISSUE VARIABILITY IN A MULTICELLUALR BODY

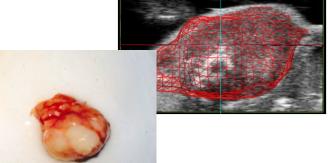




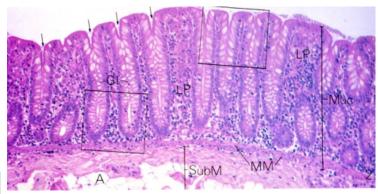


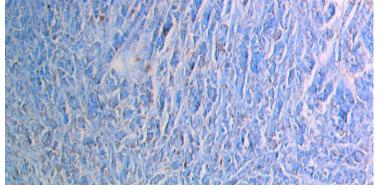








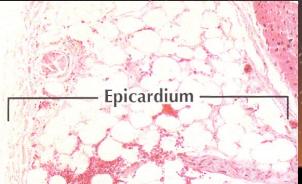




#### TISSUES AND ORGANS

- 6 × 10<sup>13</sup> CELLS of 200 different types
- cells form functional, three-dimensional, organized aggregations of morphologically similar cells and their products or derivatives TISSUES
- -- tissues constitutes **ORGANS** and organ systems







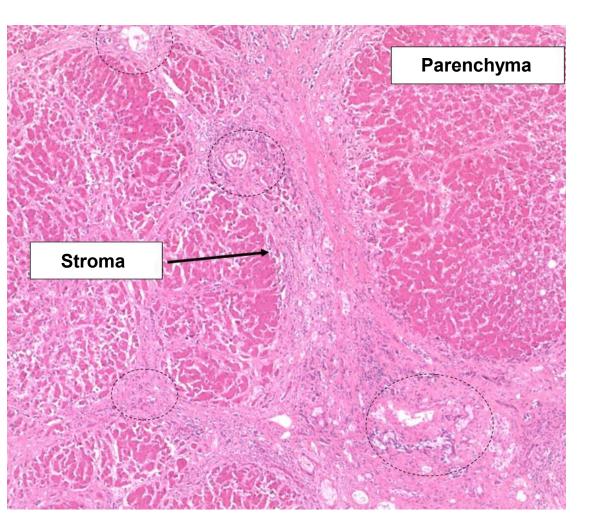


25-40×10<sup>9</sup>

#### TISSUES AND ORGANS

**Parenchyma**: functional component of a tissue (liver, lung, pancreatic, kidney parenchyma)

Stroma: surrounding, supportive tissue



#### **LIVER**

#### Parenchyma:

- Hepatocytes
- Sinusoids and adjacent structures

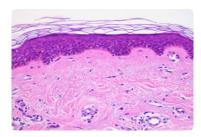
#### Stroma:

- Connective tissue and adjacent structures
- Vessels
- Nerves
- Bile ducts

#### CONTEMPORARY TISSUE CLASSIFICATION

#### Based on morphology and function:

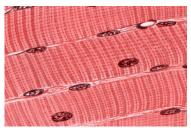
## Epithelium



Continual, avascular layers of cells with different function, oriented to open space, with specific junctions and minimum of ECM and intercellular space.

Derivates of all three germ layers

## Muscle



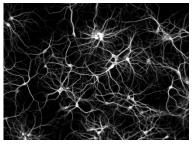
Myofibrils → contraction

Mesoderm | skeletal muscle | m

Mesoderm – skeletal muscle, myocard, mesenchyme – smooth muscles

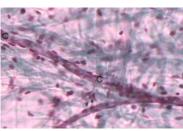
Rarely ectoderm (eg. m. sphincter a m. dilatator pupillae)

## Nerve



Neurons and neuroglia
Reception and transmission of electric signals
Ectoderm, rarely mesoderm (microglia)

## Connective

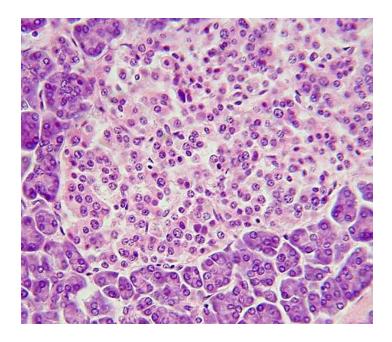


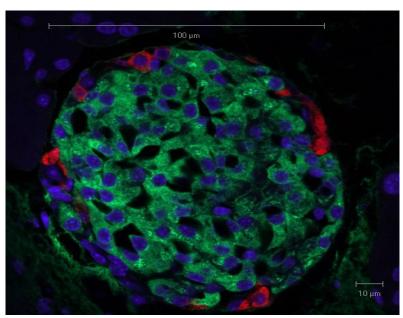
Dominant extracellular matrix Connective tissue, cartilage, bone... Mesenchyme

#### TISSUE DEFINITION

Functional, three-dimensional, organized aggregation of morphologically similar cells, their products and derivatives

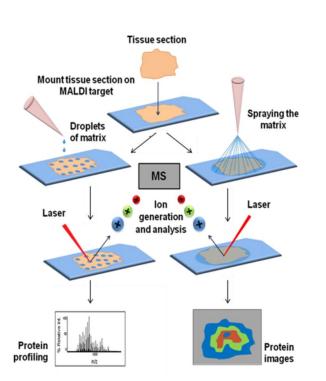
classical histological definition is based on microscopic visualization

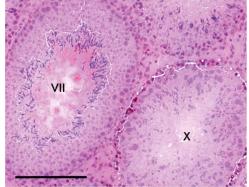


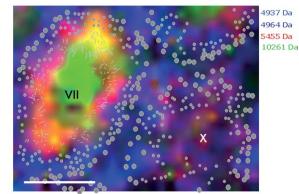


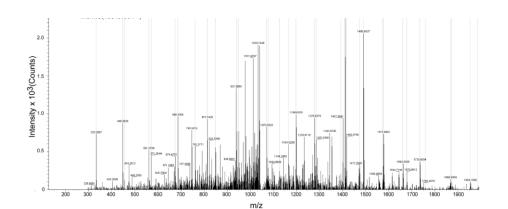
## HISTOLOGY IS NOT ONLY ABOUT MICROSCOPY











## BASIC PRINCIPLES OF HISTOGENESIS

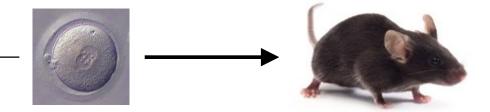
**Proliferation** 

Diferentation

Migration

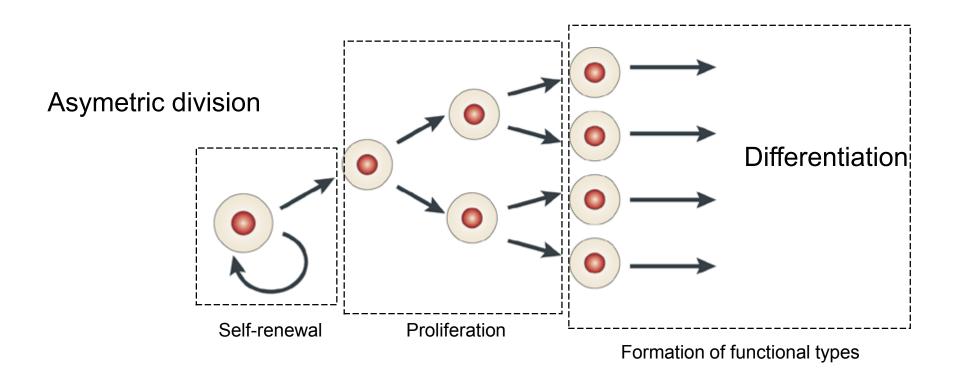
**Apoptosis** 

Tissue patterns



#### FUNCTIONAL CELL TYPES DIFFERENTIATE FROM STEM CELLS

Stem cells are capable of differentiation and self-renewal



#### STEM CELLS

#### **Totipotent**

- Constitute all cells of the body incl. extraembryonic tissues
- Zygote and early stages











#### **Pluripotent**

- All cells in the body except for trophoblast
- Blastocyst Inner cell mass ICM (embryoblast)
- Embryonic stem cells



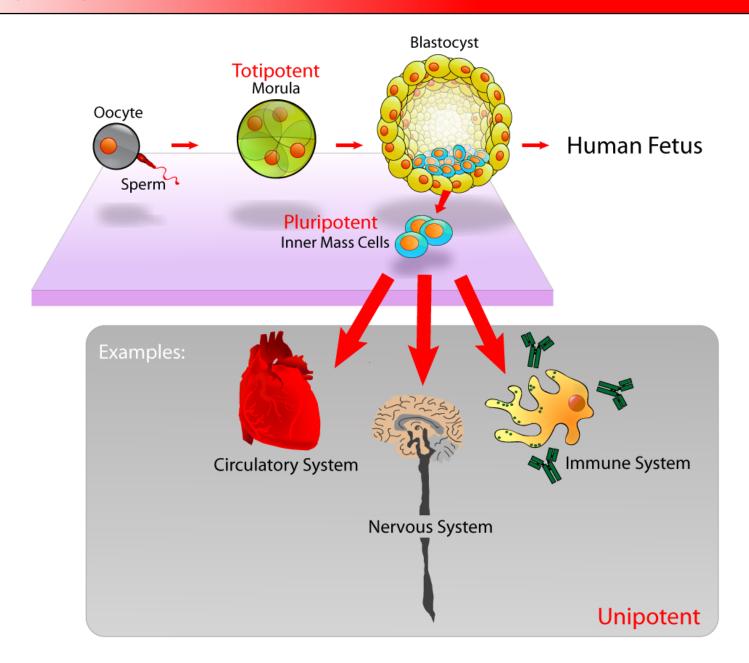
#### Multipotent

- Give rise to various cell types of a particular tissue
- Mesenchymal SC, hematopoietic SC

# Pre-T cell CLP Pre-B cell MEP Pre-B cell Megakaryocyte Pre-B cell Megakaryocyte Pre-B cell Megakaryocyte Pre-B cell Megakaryocyte Platelets Megakaryocyte Platelets Megakaryocyte Platelets Neutrophil Neutrophil Monocyte/ Macrophage

#### Oligo- a unipotent

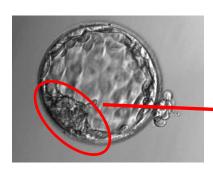
- One or several cell types – hematopoietic, tissue precursors for renewal of intestinal epithelia, etc.

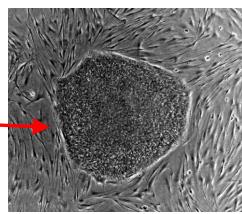


#### STEM CELLS IN HUMAN BODY

#### **Embryonic stem cells (ESCs)**

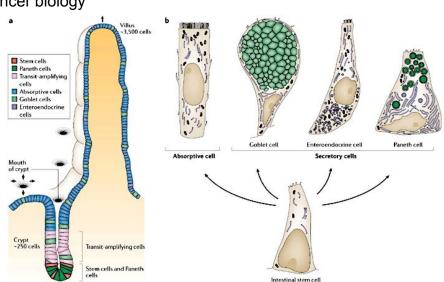
- embryoblast of blastocyst
- pluripotent
- modelling of early embryogenesis, regenerative medicine

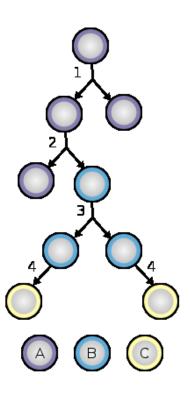




#### Tissue (adult) stem cells

- regeneration and renewal of tissues
- GIT, CNS, mesenchyme
- regenerative medicine, cancer biology





#### STEM CELLS AS RESEARCH TOOLS

#### Induced pluripotent stem cells (iPSc)

- adult differentiated cell (fibroblast) is reprogrammed into pluripotent state
- differentiation into desired cell type
- regenerative medicine, cell and gene therapy



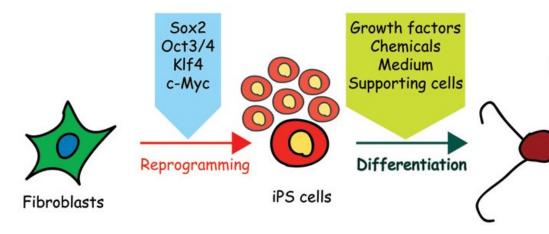




#### **Induction of Pluripotent Stem Cells** from Mouse Embryonic and Adult Fibroblast Cultures by Defined Factors

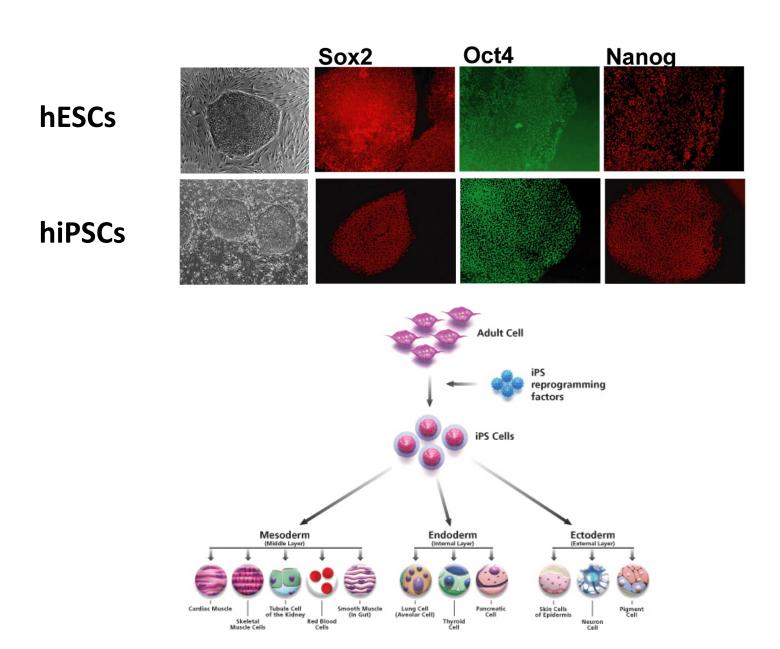
Kazutoshi Takahashi<sup>1</sup> and Shinya Yamanaka<sup>1,2,\*</sup>

Department of Stem Cell Biology, Institute for Frontier Medical Sciences, Kyoto University, Kyoto 606-8507, Japan <sup>2</sup>CREST, Japan Science and Technology Agency, Kawaguchi 332-0012, Japan \*Contact: yamanaka@frontier.kyoto-u.ac.jp DOI 10.1016/j.cell.2006.07.024



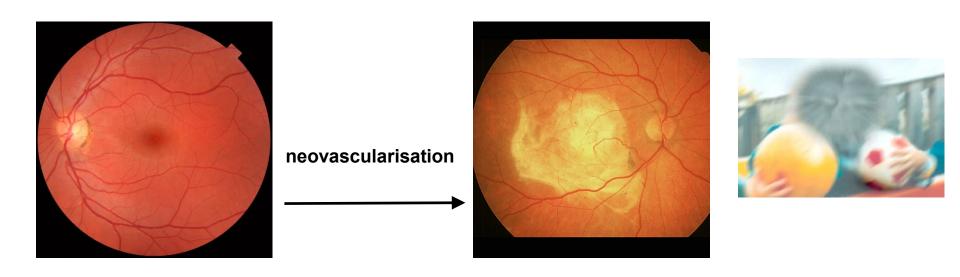
Disease modelling Drug testing Tissue replacement

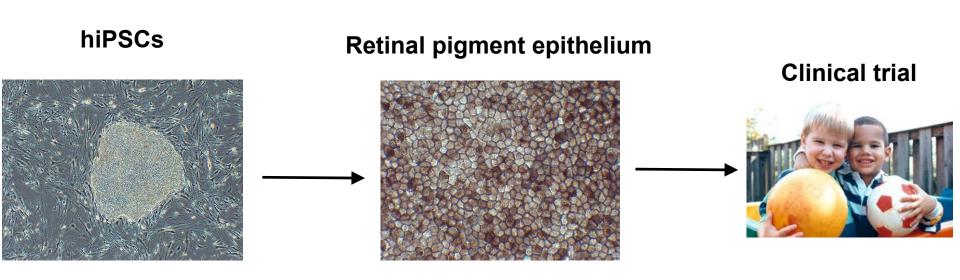
## iPSCs SHARE FUNDAMENTAL PROPERTIES WITH hESCs



## STEM CELLS AS THERAPY

## Age-related macular degeneration



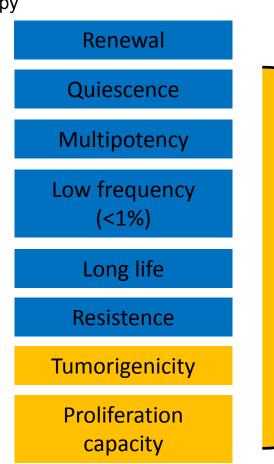


#### STEM CELLS AS FOES

#### Cancer stem cells

- solid tumor is always heterogeneous
- small population of cells with stem cell character can repopulate tumor tissue after cytotoxic therapy

Tissue stem cells



CSC-Targeted Cancer
Therapy

CSC

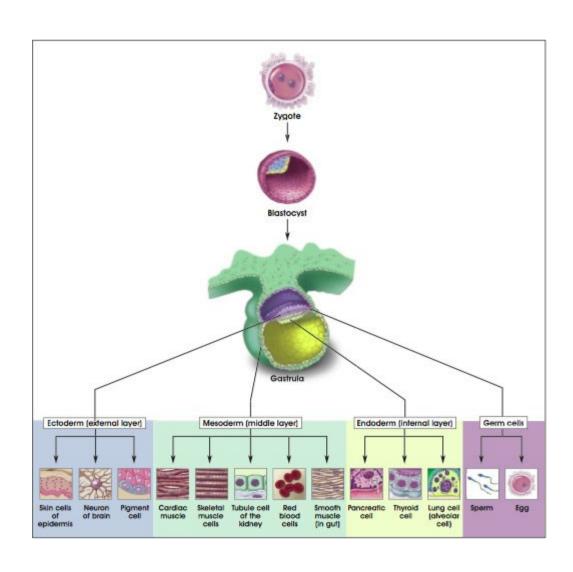
Traditional
Cancer
Therapy

Drugs that kill
tumor cells...

Drugs that kill
tumor cells...

Tumor shrinks,
but grows back

Cancer stem cells



#### **CELL DIFFERENTIATION**

Induction of differentiation

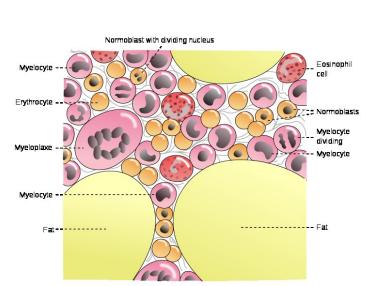
-blast

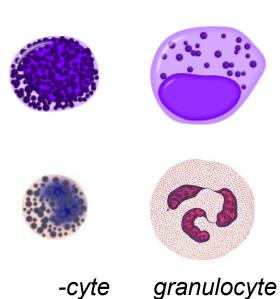
**Determination and commitment** 



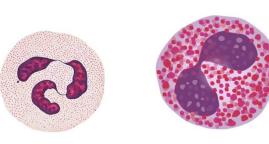
eg. myeloblast

**Terminal differentiation** 

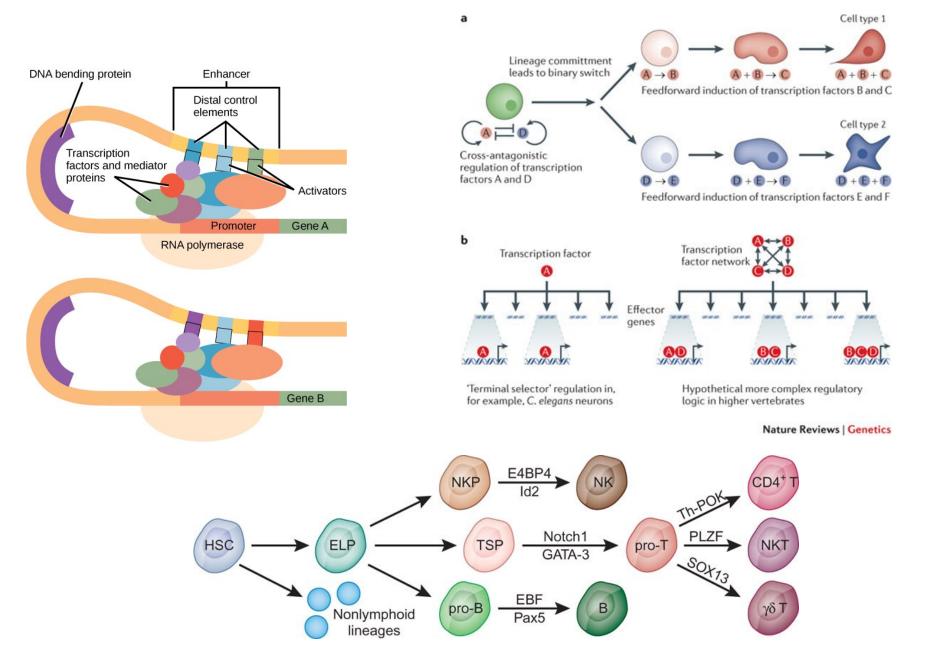




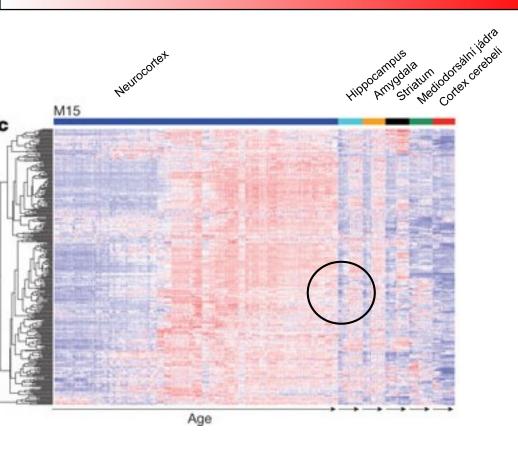


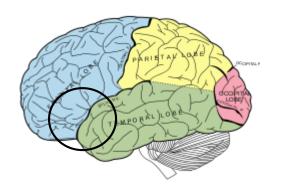


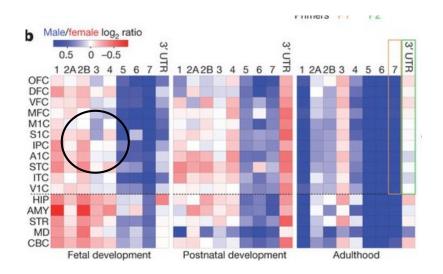
#### DIFFERENTIATION IS DRIVEN BY GENE TRANSCRIPTION



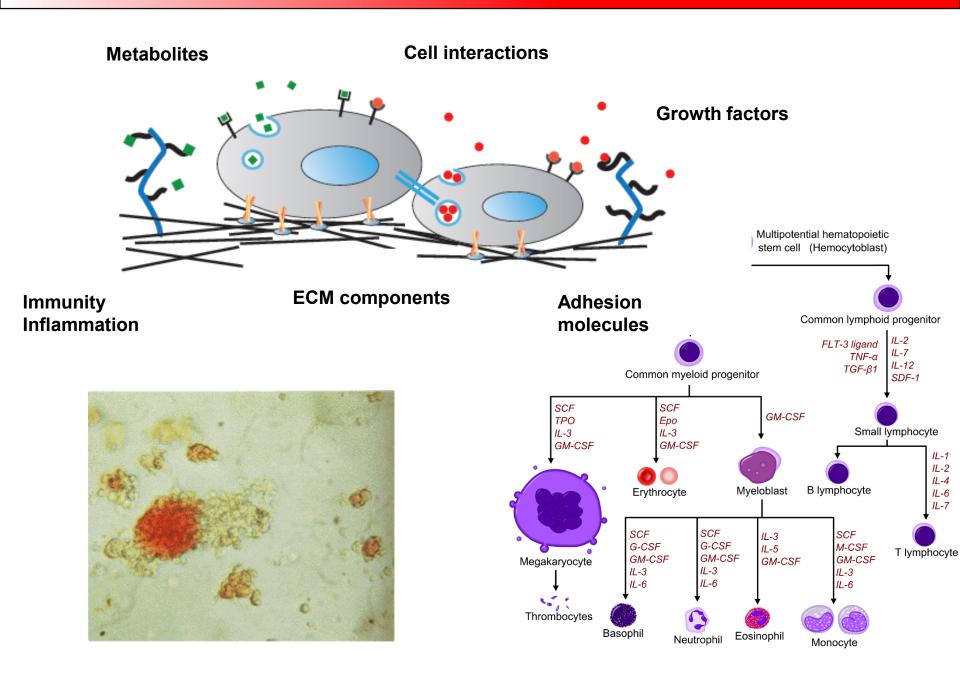
#### TISSUE DIFFER IN THEIR GENETIC AND EPIGENETIC PROFILES







#### CELLS CAN CREATE UNIQUE MICROENVIRONMENT



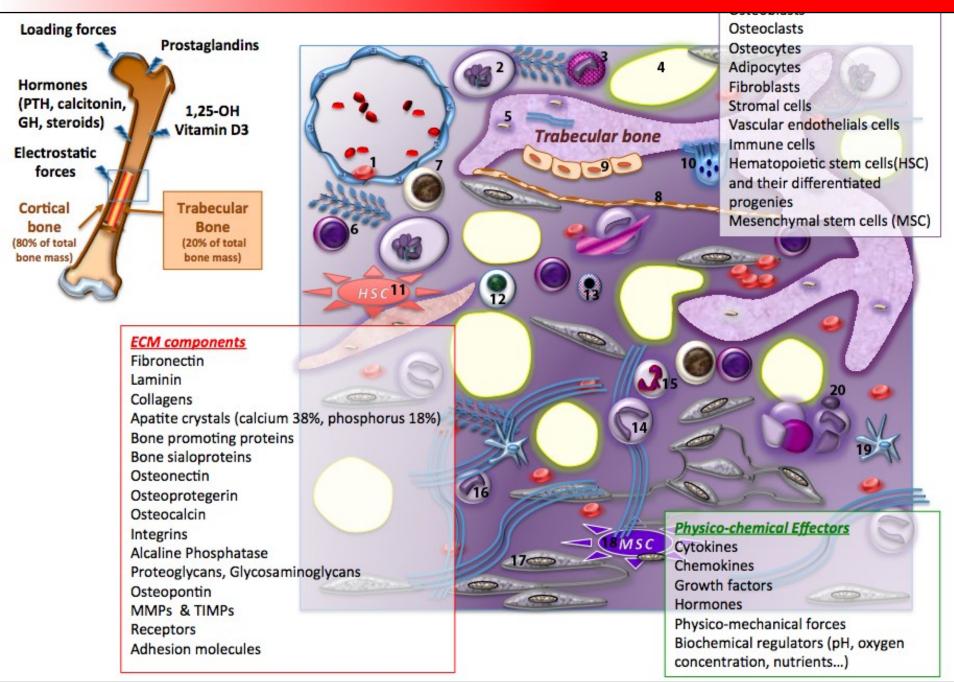
#### MICROENVIRONMENT REGULATES TISSUE FUNCTION

Huge number of biological and physically-chemical parameters

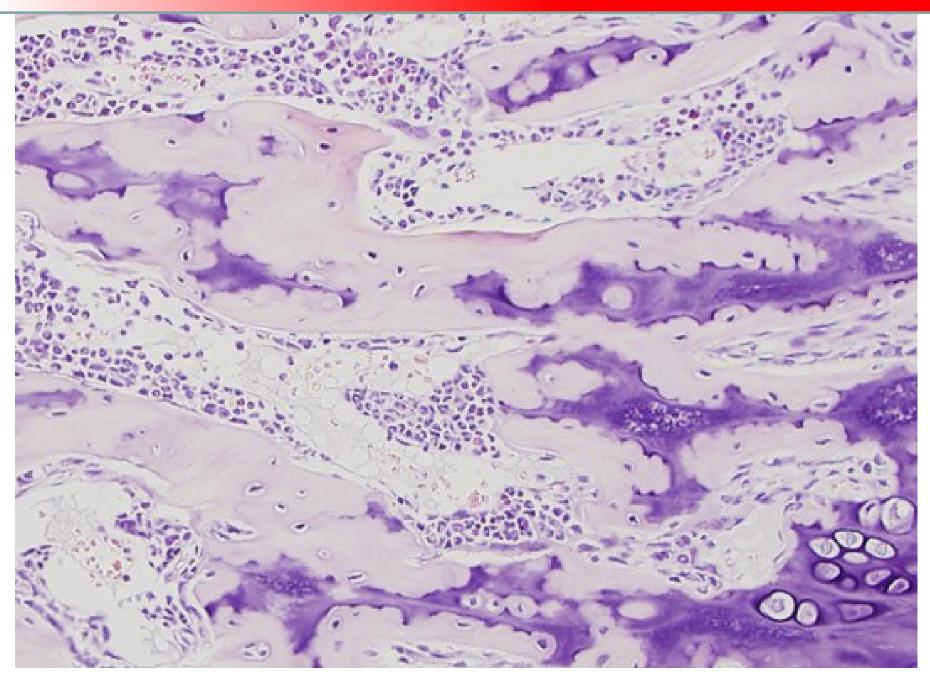


- Embryonic development
- Intercellular interaction
- Space organization (dimensionality)
- Gradient of morphogenes
- Epigenetic profile
- Gene expression dynamics
- Partial pressure of gases
- ECM composition
- Mechanical stimulation
- Perfusion and interstitial flows
- Local immunity response
- Metabolites

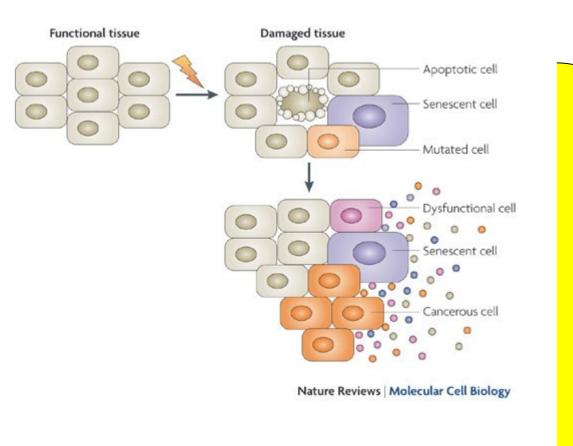
#### STEM CELL NICHE



## HEMATOPOIETIC NICHE



#### MICROENVIRONMENT IS NECESSARY FOR TISSUE HOMEOSTASIS



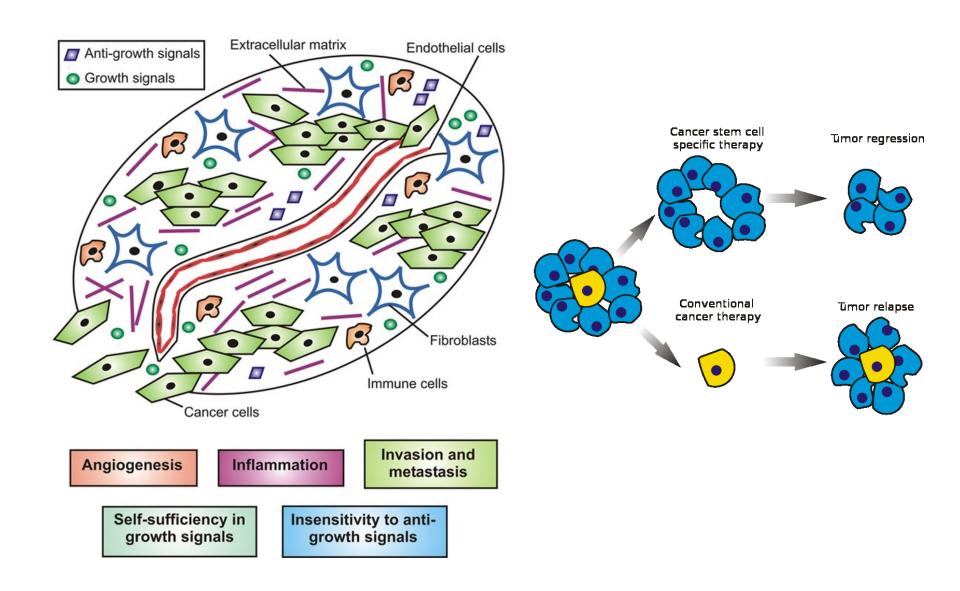
**Apoptosis** 

Regeneration

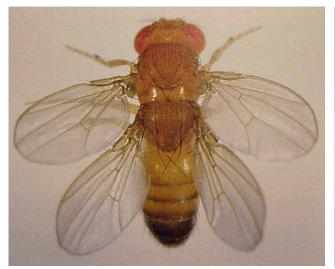
Senescence

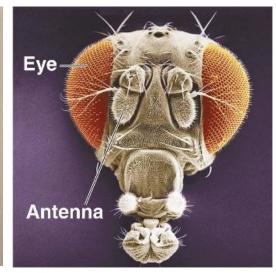
**Transformation** 

#### MICROENVIRONMENT IS OF CLINICAL IMPORTANCE



## MOLECULAR PRINCIPLES OF HISTOGENESIS





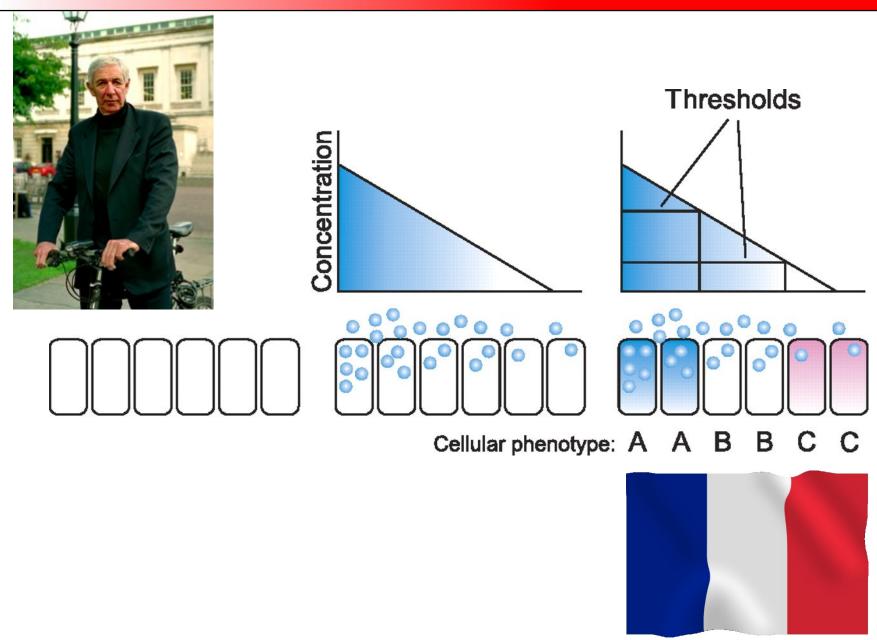


Wild type

Mutant



#### LEWIS WOLPERT AND FRENCH FLAG MODEL

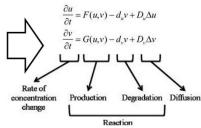


#### WHY DO TIGERS HAVE STRIPES?

#### Reakčně-difúzní systém

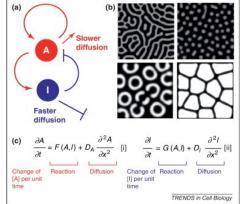












#### THE CHEMICAL BASIS OF MORPHOGENESIS

By A. M. TURING, F.R.S. University of Manchester

(Received 9 November 1951—Revised 15 March 1952)

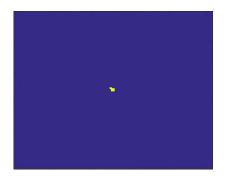
It is suggested that a system of chemical substances, called morphogens, reacting together and diffusing through a tissue, is adequate to account for the main phenomena of morphogenesis. Such a system, although it may originally be quite homogeneous, may later develop a pattern or structure due to an instability of the homogeneous equilibrium, which is triggered off by random disturbances. Such reaction-diffusion systems are considered in some detail in the case of an isolated ring of cells, a mathematically convenient, though biologically unusual system. The investigation is chiefly concerned with the onset of instability. It is found that there are six essentially different forms which this may take. In the most interesting form stationary waves appear on the ring. It is suggested that this might account, for instance, for the tentacle patterns on Hydra and for whorled leaves. A system of reactions and diffusion on a sphere is also considered. Such a system appears to account for gastrulation. Another reaction system in two dimensions gives rise to patterns reminiscent of dappling. It is also suggested that stationary waves in two dimensions could account for the phenomena of phyllotaxis.

The purpose of this paper is to discuss a possible mechanism by which the genes of a zygote may determine the anatomical structure of the resulting organism. The theory does not make any new hypotheses; it merely suggests that certain well-known physical laws are sufficient to account for many of the facts. The full understanding of the paper requires a good knowledge of mathematics, some biology, and some elementary chemistry. Since readers cannot be expected to be experts in all of these subjects, a number of elementary facts are explained, which can be found in text-books, but whose omission would make the paper difficult reading.

#### 1. A model of the embryo. Morphogens

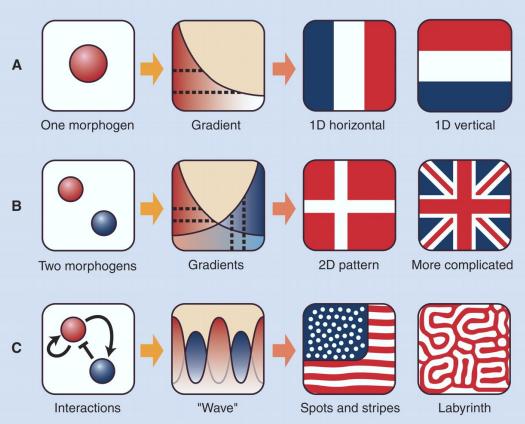
In this section a mathematical model of the growing embryo will be described. This model will be a simplification and an idealization, and consequently a falsification. It is to be hoped that the features retained for discussion are those of greatest importance in the present state of knowledge.

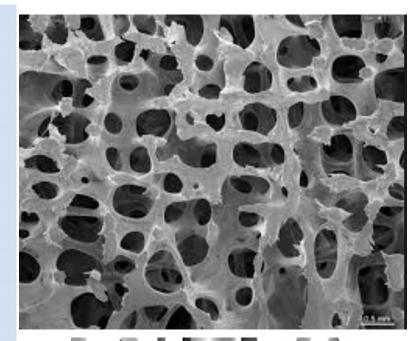
The model takes two slightly different forms. In one of them the cell theory is recognized but the cells are idealized into geometrical points. In the other the matter of the organism is imagined as continuously distributed. The cells are not, however, completely ignored, for various physical and physico-chemical characteristics of the matter as a whole are assumed to have values appropriate to the cellular matter.



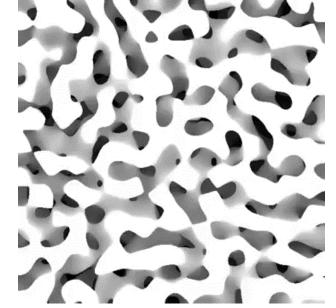


#### TISSUE PATTERNS ARE DRIVEN BY GRADIENTS OF MORPHOGENES

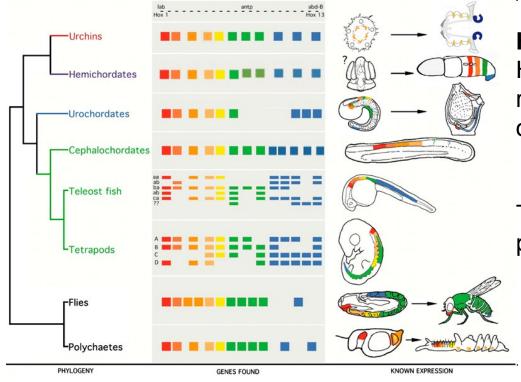








#### HOX COMPLEX

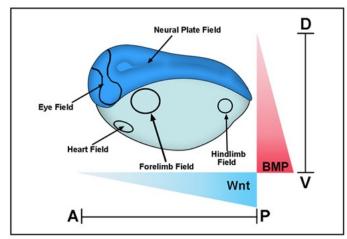


#### Hox genes

Highly conserved family of transcription regulators that determine body polarity, orientation and axis

Tissue differentiation along anterioposterior axis

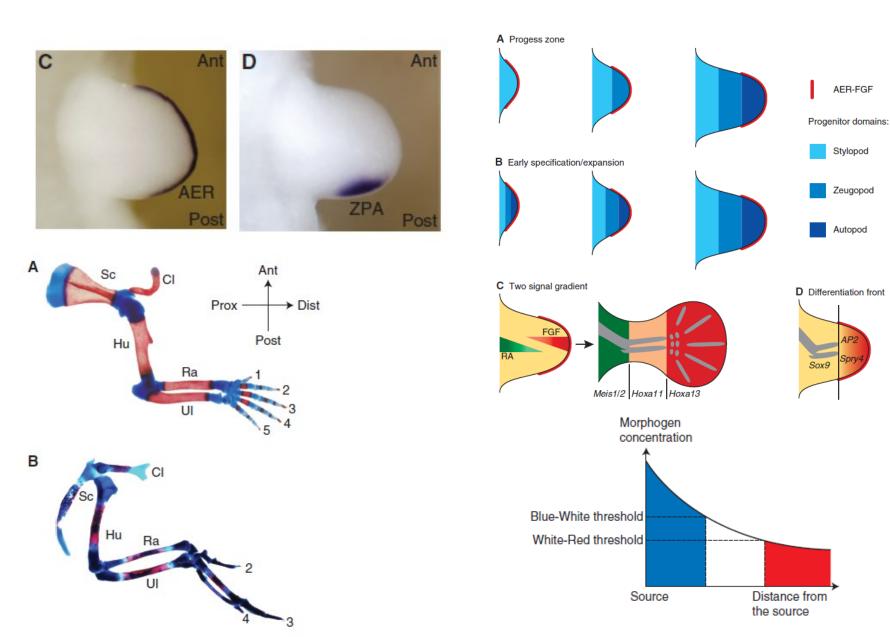
doi:10.1038/sj.hdy.6800872



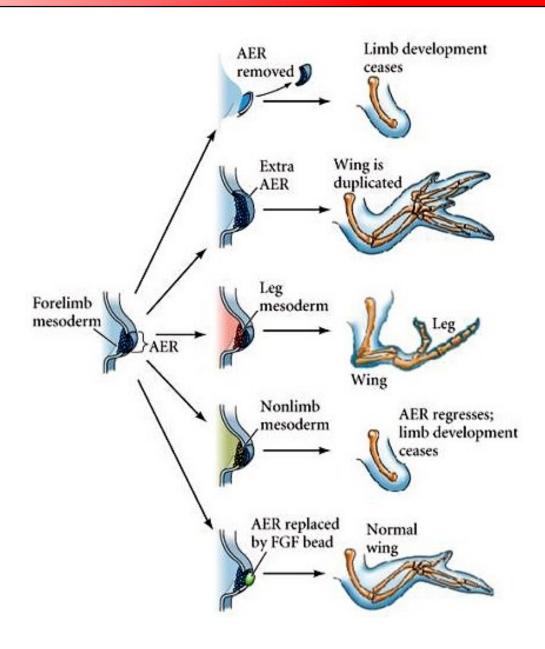
Human (39 genes)

Cluster	Chromosome	# Hox genes
HoxA	7	11
HoxB	17	10
HoxC	12	9
HoxD	2	9

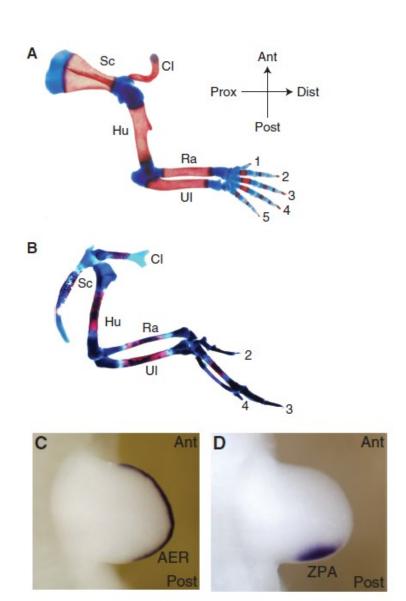
# TEMPORO-SPATIAL EXPRESSSION OF MORPHOGENES DRIVES FINAL LOCALIZATION, ORIENTATION AND MORPHOLOGY OF TISSUES AND ORGANS

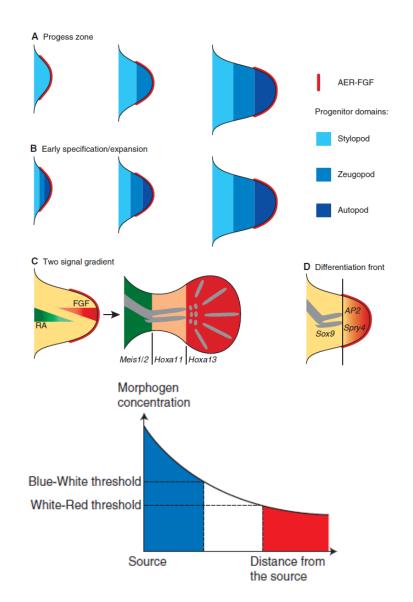


#### MANIPULATING AER ALTERS INSTRUCTIONS FOR LIMB DEVELOPMENT

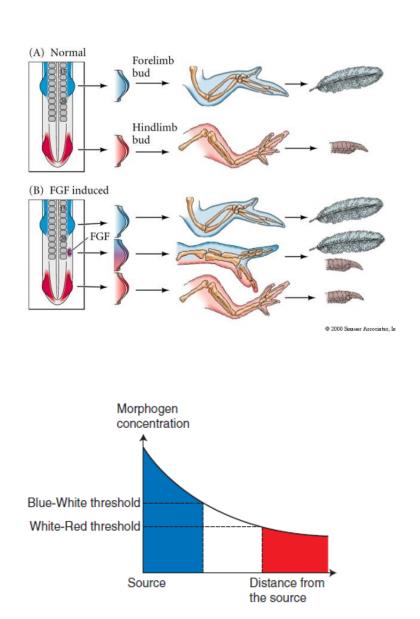


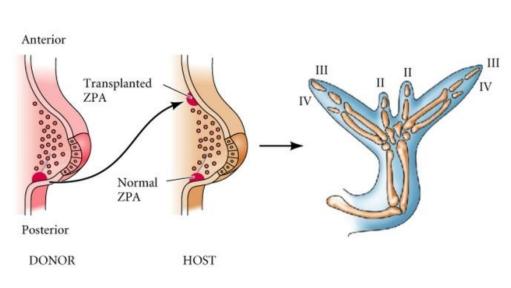
## TEMPORO-SPATIAL EXPRESSION OF DIFFERENT REGULATORS DETERMINES FINAL LOCALIZATION, ORIENTATION AND MORPHOLOGY OF TISSUES AND ORGANS





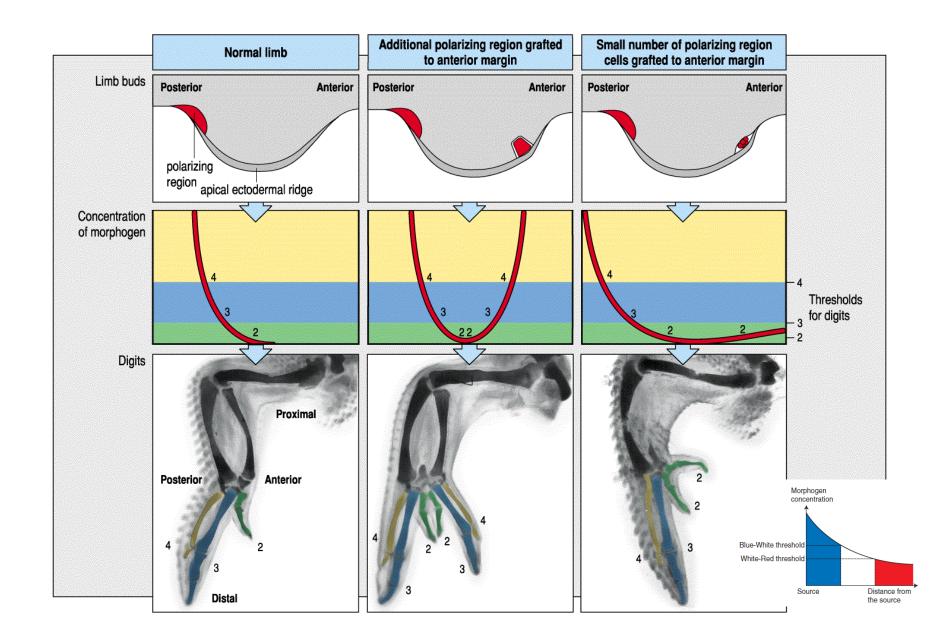
## ZPA SPECIFIES POSITIONAL INFORMATION IN LIMB BUD



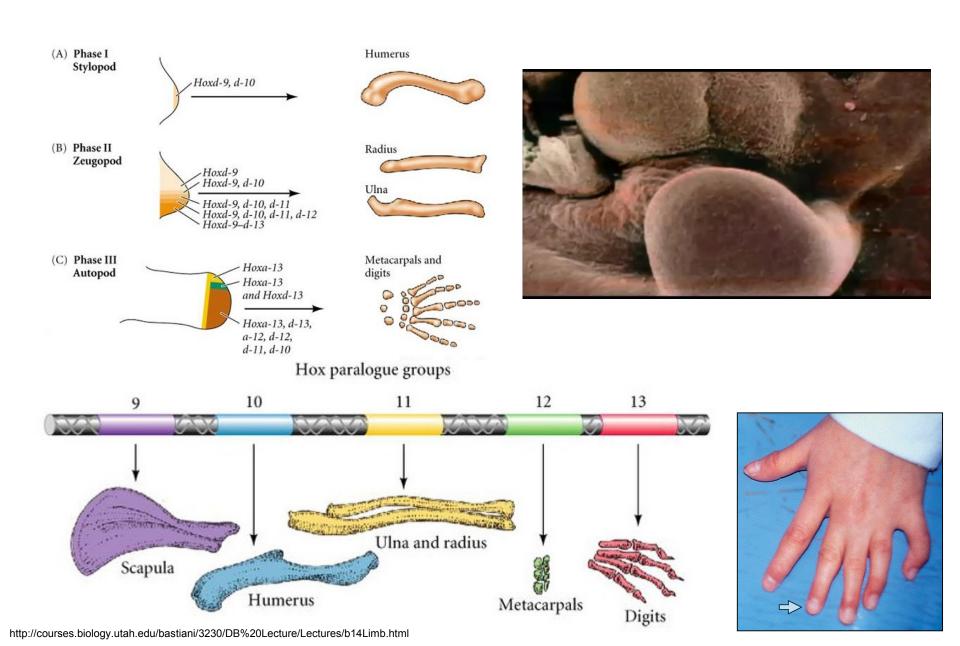


@ 2000 Sinauer Associates, Inc.

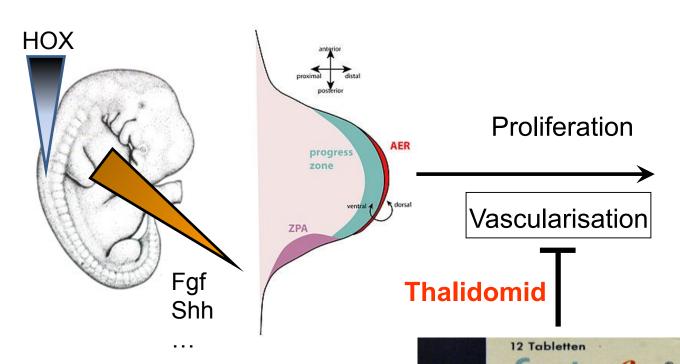
## MORPHOGENES FROM AER AND ZPA DEFINES LIMB FORMATION



## HOX PATTERN DRIVES TRANSCRIPTIONAL RESPONSE



## STORY OF THALIDOMID

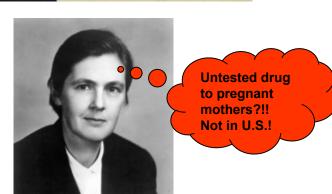


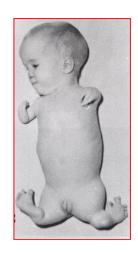
GRUNENTHAL



## Thalidomid embryopathy

- phocomelia
- amelia
- anophtalmia/microphtalmia
- abnormal kidneys, heart, GIT, genitalia

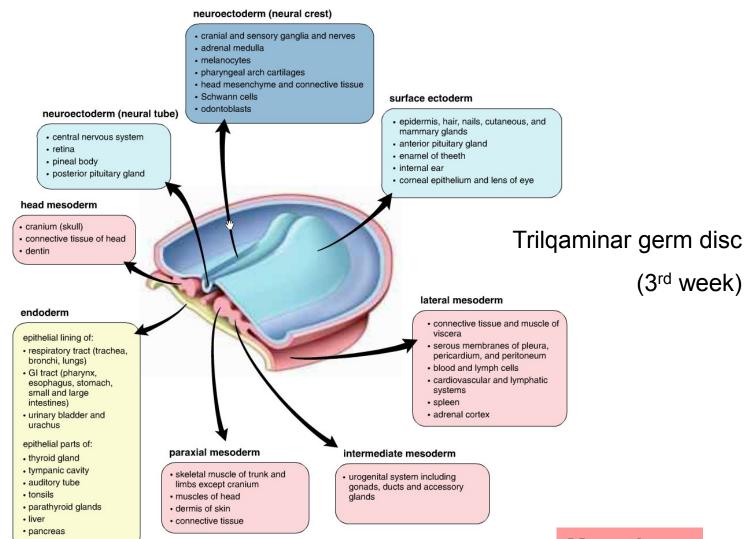




Frances Oldham Kelsey, FDA USA

## HISTOGENESIS AND ORGANOGENESIS

## **Ectoderm**



**Endoderm** 

Mesoderm

#### **Ectoderm**

- Epidermis, hair nails, cutaneous and mammary glands
- Corneal epithelium and lens of eye
- Enamel of teeth
- Internal ear
- Anterior pituitary gland
- Epithelium of oral cavity and part of anal canal
- Neural tube and derivatives
- CNS
- Retina
- Posterior pituitary gland
- Pineal body
- Neural crest and derivatives:
- Cranial and sensory ganglia and nerves
- Schwann cells
- adrenal medulla
- Enteroendocrinne cells
- Melanocytes
- Head mesenchyme and connective tissue
- Odontoblasts

#### Mesoderm

#### Connective tissue of head

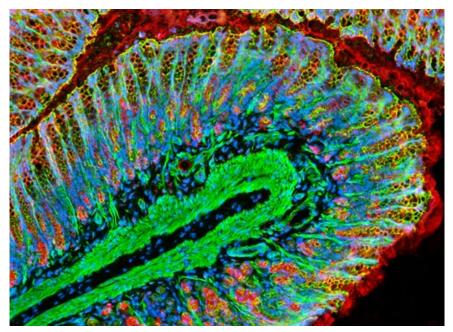
- Cranium, dentin
- Skeletal muscle of trunk and limbs Paraxial except cranium
  - Dermis of skin

Intermediate

- Muscles of head
- Urogenital system + ducts, glands and gonads

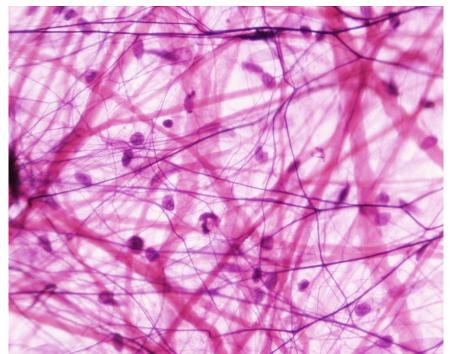
#### **Endoderm**

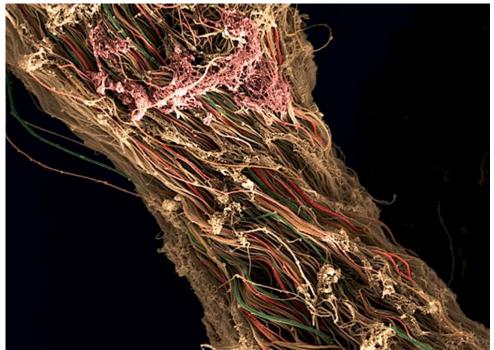
- GIT epithelium except oral cavity and part of anal canal
- Extramural glands of GIT
- Epithelium of bladder
- Epithelium of respiratory system
- Thyroid gland, parathyroid glands, thymus
- Tonsils
- Epithelium of cavum tympani and Eustachian tube
- Visceral muscle and connective tissue
- Serous membranes of pleura, peritoneum and pericardium
- Blood cells, leukocytes
- Cardiovascular and lymphatic system
- Spleen
- Adrenal cortex



## 6. CONNECTIVE TISSUE

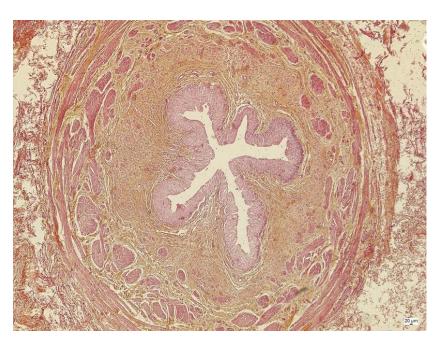
Not just a tissue glue...

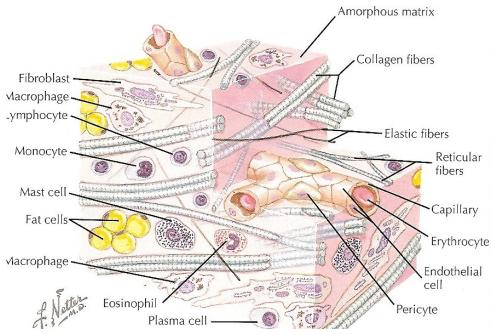




## Mechanical and biological properties

→ surrounds other tissues, compartmentalization, support, physico-chemical environment, immunological support, storage





## GENERAL COMPOSITION OF CONNECTIVE TISSUE

#### Cells and extracellular matrix

Cells

**Connective tissue** – permanent and transient cell populations (fibroblasts/myofibroblasts,

immune cells, adipocytes, adult stem cells)

Cartilage – chondroblasts/chondrocytes

**Bone** – osteoblasts/osteocytes/osteoclasts

Matrix – fibrous and amorphous

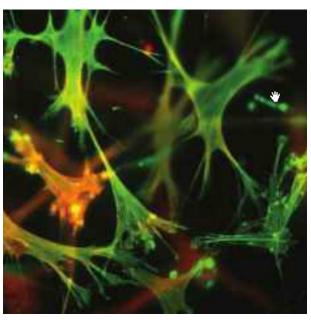
#### Fibrous component

- collagen
- reticular
- elastic

**Amorphous component** (amorphous ground substance)

- Complex matrix consisting of glycosaminoglycans, glycoproteins and proteoglycans,

depending on tissue type (connective  $\times$  ligament  $\times$  cartilage  $\times$  bone)



#### CLASSSIFICATION OF CONNECTIVE TISSUE

## **Embryonic CT**

- Mesenchyme
- Jelly-like CT (Wharton jelly, dental pulp, stroma of iris)

#### Adult CT

- Areolar (loose, interstitial) CT
- Dense collagen irregular CT
- Dense collagen regular CT
- Fat (adipose tissue)
- Cartilage
- Bone
- Blood and hematopoietic tissue
- Lymphatic tissue

CT

Specialized CT

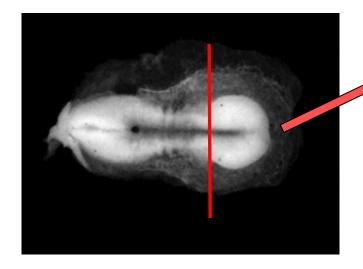
Trophic CT (body liquids)

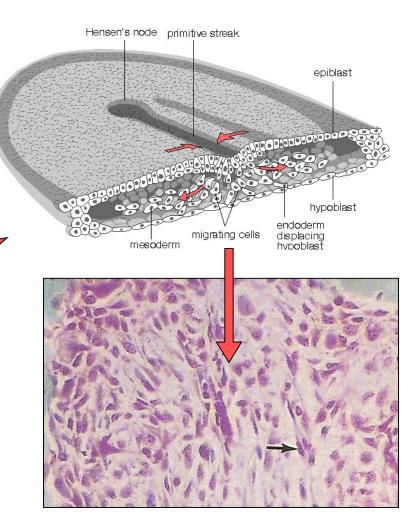
## EMBRYONIC ORIGIN OF CONNECTIVE TISSUE

- Mesenchyme = loose tissue between germ layers
- Complex network of star- or spindle-shaped cells

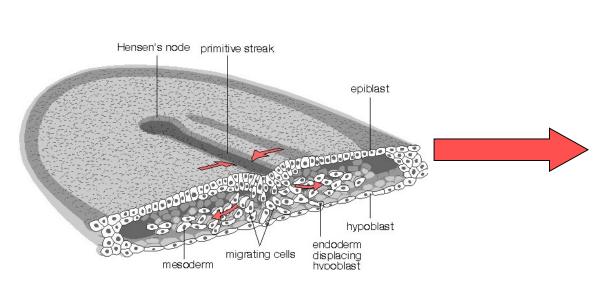
Jelly-like amorphous ground substance

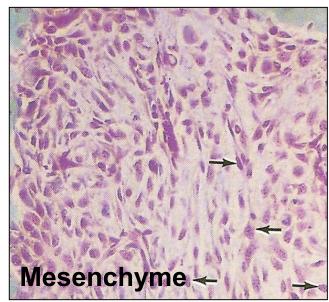
DAY 12 of embryonic development

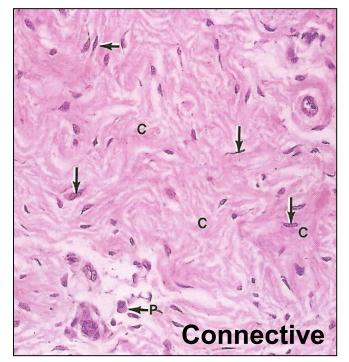


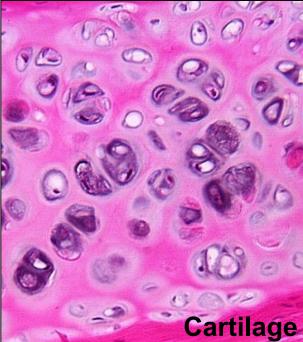


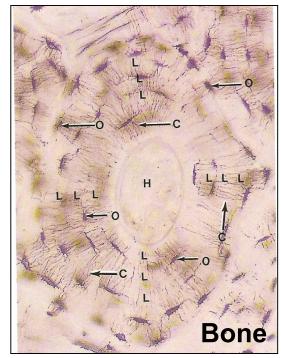
## DERIVATIVES OF CONNECTIVE TISSUE











## **CELLS OF CONNECTIVE TISSUE**

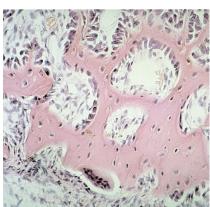
#### Cells

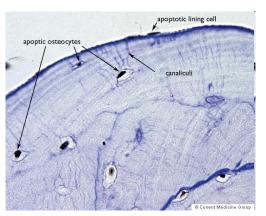
- Fibroblasts/fibrocytes/myofibroblasts
- Heparinocytes
- Macrophages of CT = histiocytes
- Plasma cells
- Lymphocytes
- Adipocytes
- Adult stem cells

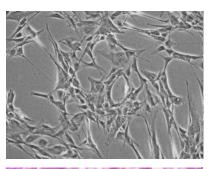
#### **Extracellular matrix**

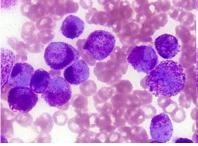
- Fibrous compound
- Amorphous ground substance

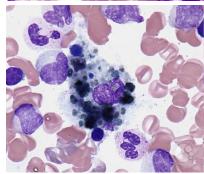


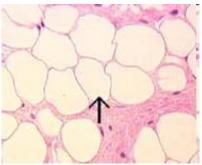






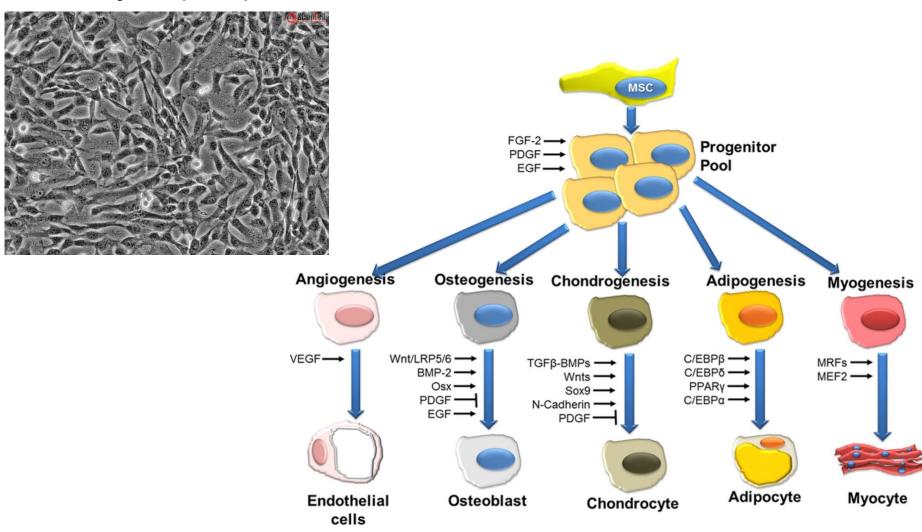






## **CELLS OF CONNECTIVE TISSUE**

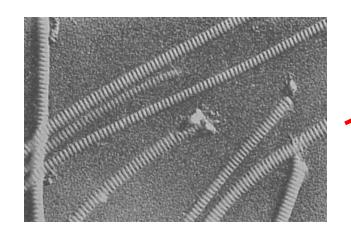
## Mesenchymal (adult) stem cells

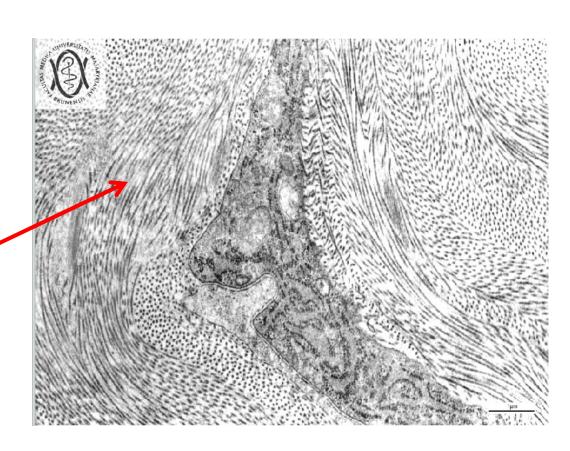


## EXTRACELLULAR MATRIX – FIBROUS COMPONENT

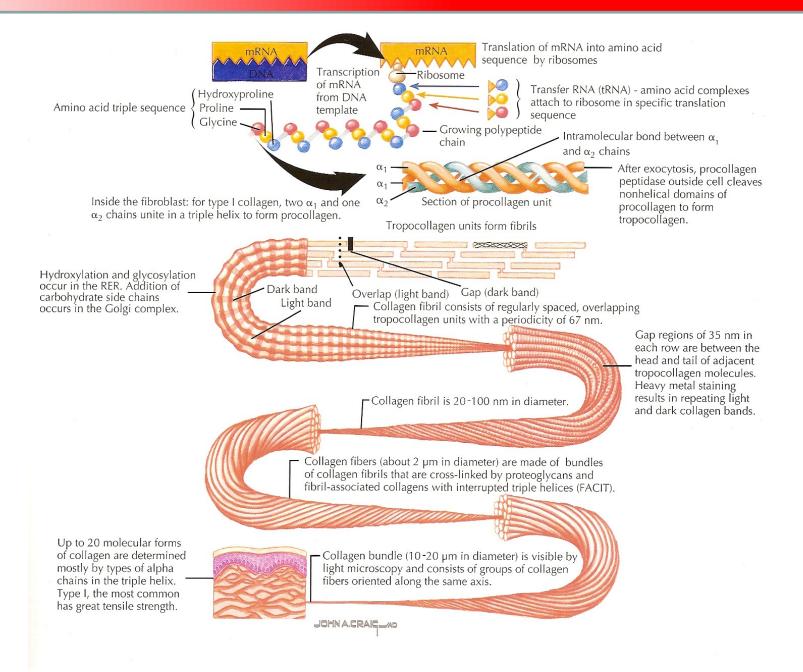
## Collagen fibers

- family of fibrous proteins encoded by >35 genes (2013)
- polymer subunit = tropocollagen; triple helix
- different structural and mechanical properties (strength, elasticity, pliability...)
- most abundant protein in human body ( 30% dry weight)





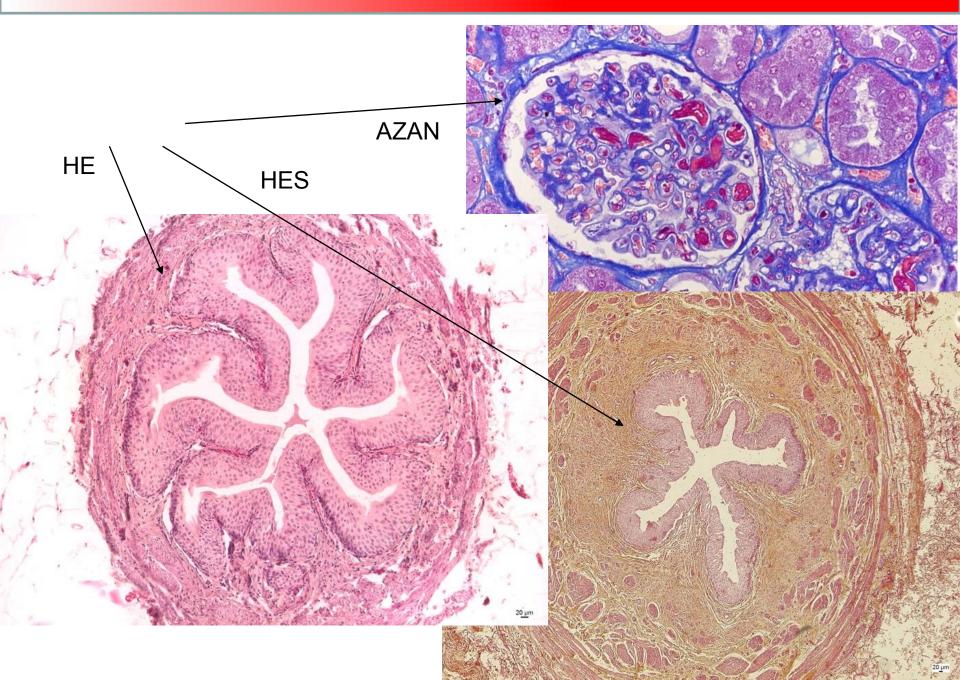
#### COLLAGEN



## COLLAGEN

Type	Localization	Structure	Main function
1	Bone, tendons, meniscus, dentin, dermis, capsules of organs, loose CT 90% of type I	Fibrils (75nm) – fibers (1-20μm)	Resilience in pull
II	Hyaline and elastic cartilage	Fibrils (20nm)	Resilience in pressure
III	Skin, veins, smooth muscles, uterus, liver, spleen, kidney, lung	Like I, high content of proteoglycans and glycoprotiens, reticular network	Shape formation
IV	Basal lamina of epithelium and endtohelium, basal membranes	No fibrils or fibers	Mechanical support
V	Lamina of muscle cells and adipocytes, fetal membranes	Like IV	
VI	Interstitial tissue, chondrocytes – adhesion		Connecting dermis and epidermis
VII	Basal membrane of epithelium		
VIII	Some endothelia (Cornea)		
Х	Growth plate, mineralized cartilage		Growth of bones, mineralization

## COLLAGEN IN LIGHT MICROSCOPE



## **COLLAGEN IN ART**

# Julian Voss-Andreae "Unraveling Collagen"

## 2005

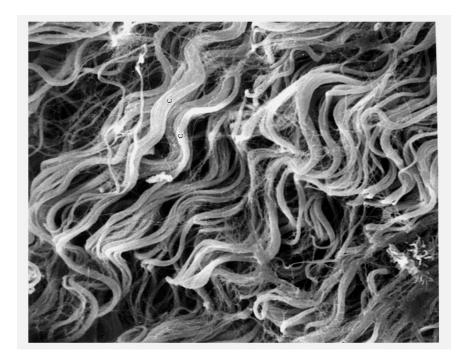
Orange Memorial Park Sculpture Garden, City of South San Francisco, CA





## **ELASTIC FIBERS**

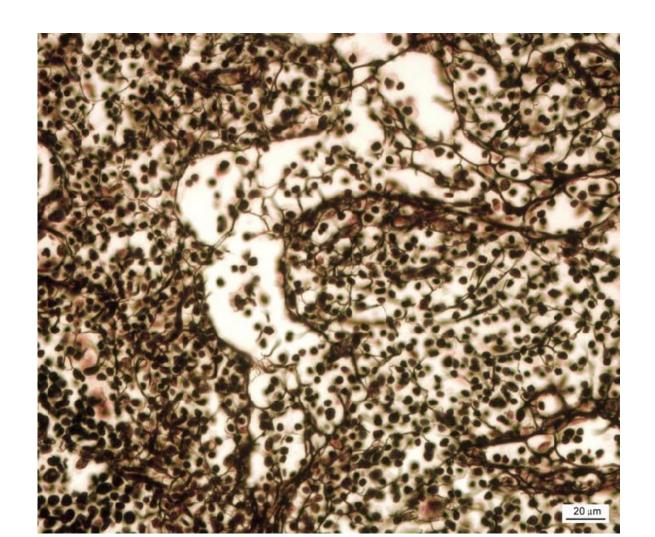
- less abundant than collagen
- polymer tropoelastin
- minimal tensile resistance, loss of elasticity if overstretched
- reduction of hysteresis = allow return back to original state after mechanic change





## RETICULAR FIBERS

- collagen 3D meshwork
- bone marrow, spleen, lymphatic nodules
- microenvironment for e.g. hematopoietic stem cells and progenitors



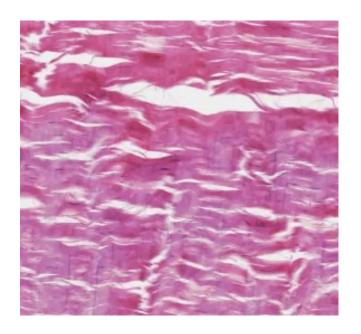
## RETICULAR CONNECTIVE TISSUE

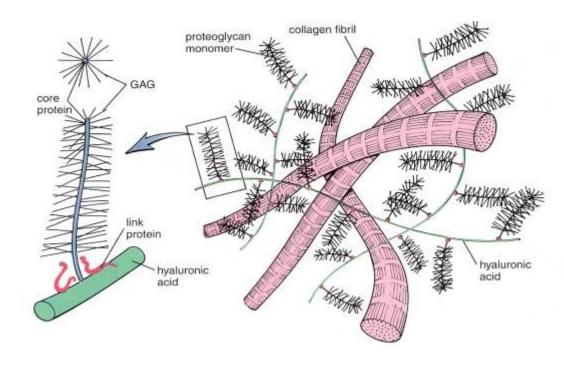


## EXTRACELLULAR MATRIX - GROUND SUBSTANCE

Amorphous extracellular matrix

Colorless, transparent, homogenous substance consisting of <u>glycosaminglycans</u>, <u>proteoglycans</u> and <u>structural glycoproteins</u>





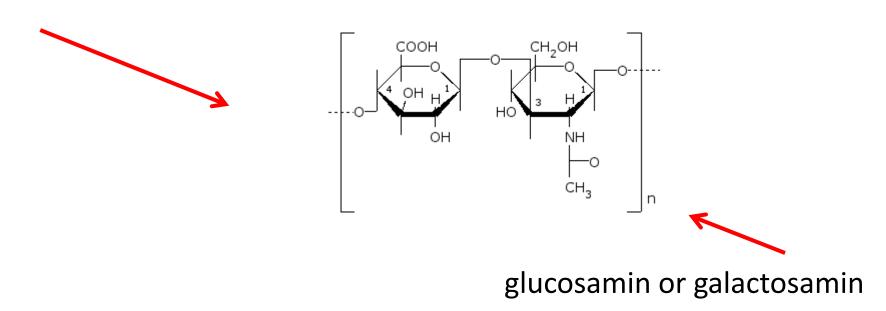
## **GLYCOSAMINOGLYCANS**

linear polysaccharides composed of two disaccharide subunits

- uronic acid and hexosamine

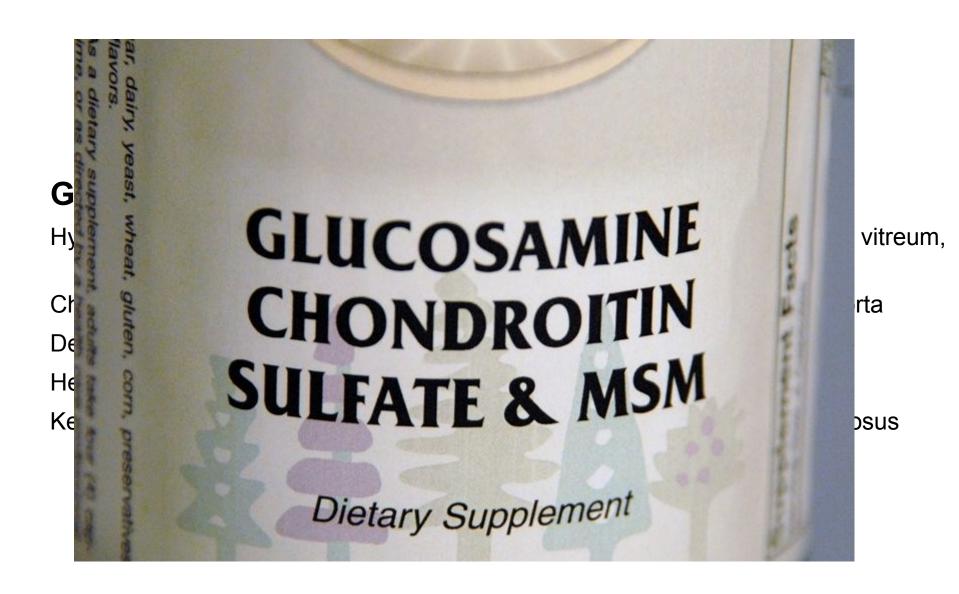
polysaccharides rich in hexosamines = acid mukopolysaccharides

## glucuronic or iduronic acid



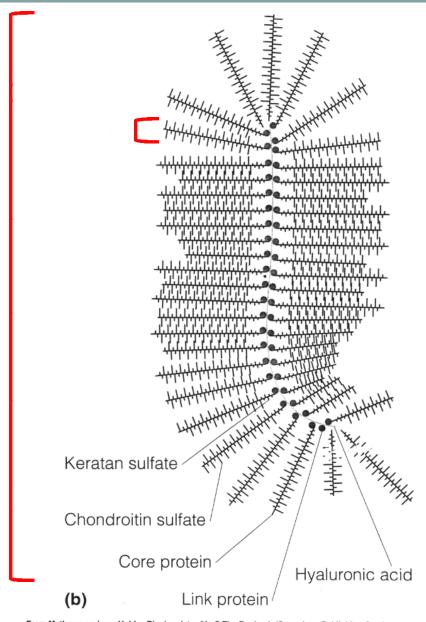
## **GLYCOSAMINOGLYCANS**

They bind to protein structures (except for hyaluronic acid)



## **PROTEOGLYCANS**

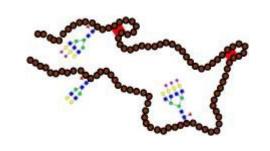
- protein + dominant <u>linear</u> saccharide component
- proteoglycan aggregates
- water-binding, volume dependent of hydratation
- aggrecan (cartilage)
- syndecan
- fibroglycan



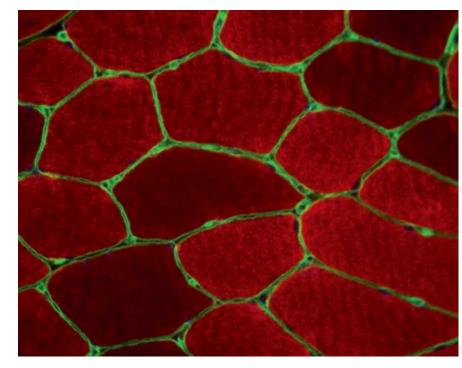
From Mathews and van Holde: Biochemistry 2/e. © The Benjamin/Cummings Publishing Co., Inc.

## STRUCTURAL GLYCOPROTEINS

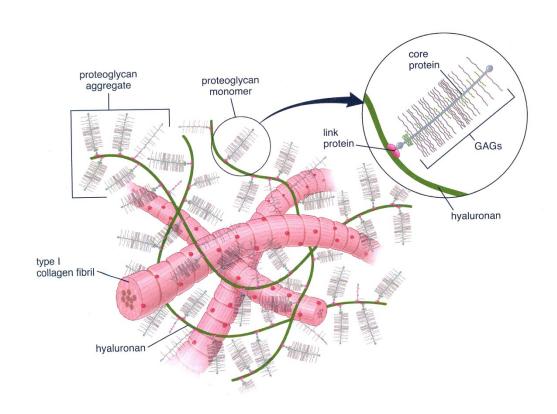
- dominant protein + <u>branched saccharide component</u>
- interaction between cells and ECM

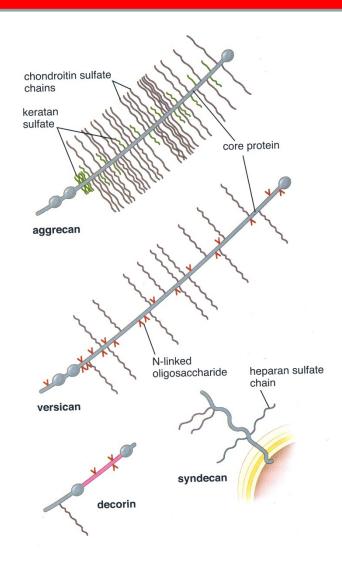


- fibronectin connects collagen fibers and glykosaminoglycans, cell adhesion and migration
- laminin basal lamina epithelial integrity
- chondronectin cartilage adhesion of chondrocytes to collagen



## **COMPOSITION OF ECM**





## CLASSIFICATION OF SPECIALIZED CONNECTIVE TISSUE

Dense
Connective Tissue
(Connective Tissue)

Compact Bone
(Connective Tissue)

Compact Bone
(Connective Tissue)

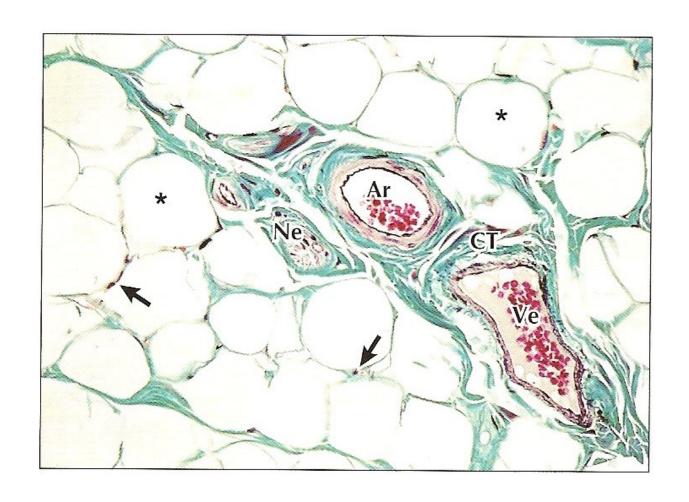
Connective Tissue)

Compact Bone
(Connective Tissue)

http://www.exploringnature.org/db/detail.php?dbID=21&detID=691

## ADIPOSE TISSUE

- Adipocytes, fibroblasts, reticular, collagen and elastic fibers, capillaries
- White and brown adipose tissue



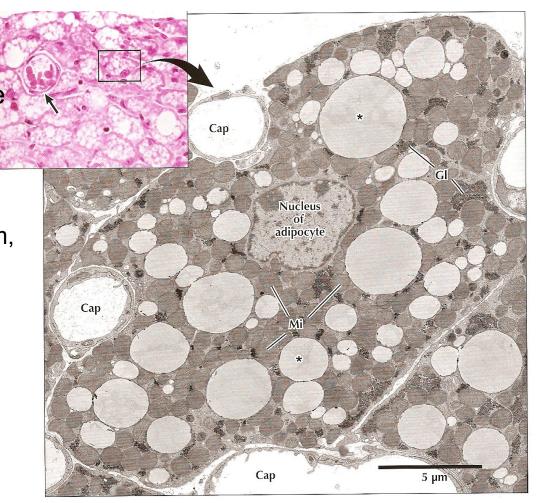
## **BROWN ADIPOSE TISSUE**

fetus and child to 1<sup>st</sup> year of life

fast source of energy

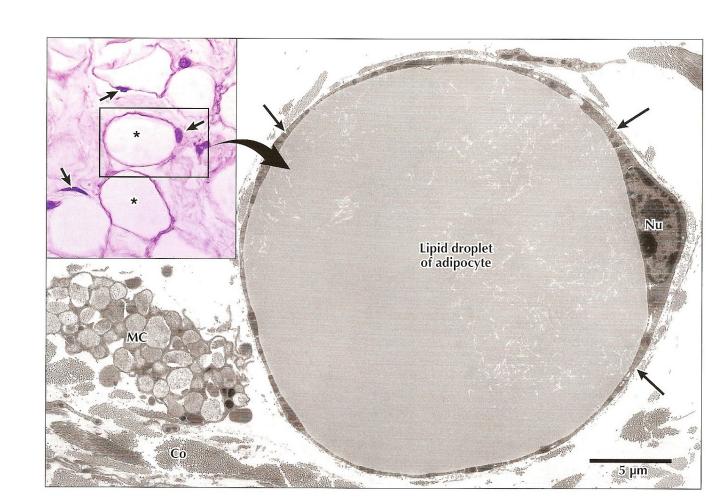
 typical localization – between shoulder blades, axilla, mediastinum, around kidneys, pancreas, small intestine

small cells with numerous fat droplets

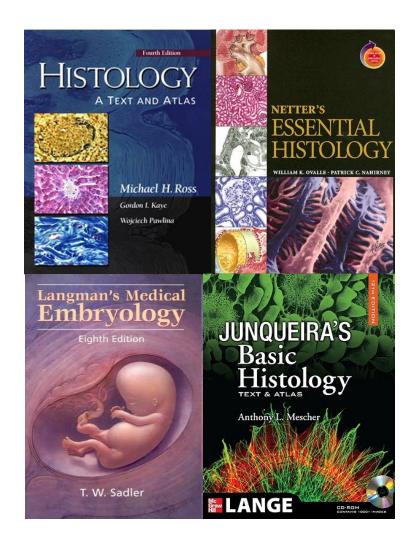


## WHITE ADIPOSE TISSUE

- adipocytes are actively formed until 2<sup>nd</sup> year of life
- no innervations, but rich vascularisation
- adipocytes with only one lipid droplet
- leptin (adipokinins)



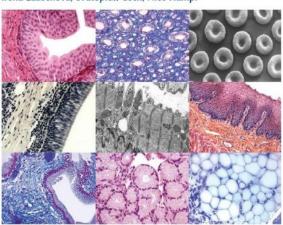
## **FURTHER STUDY**





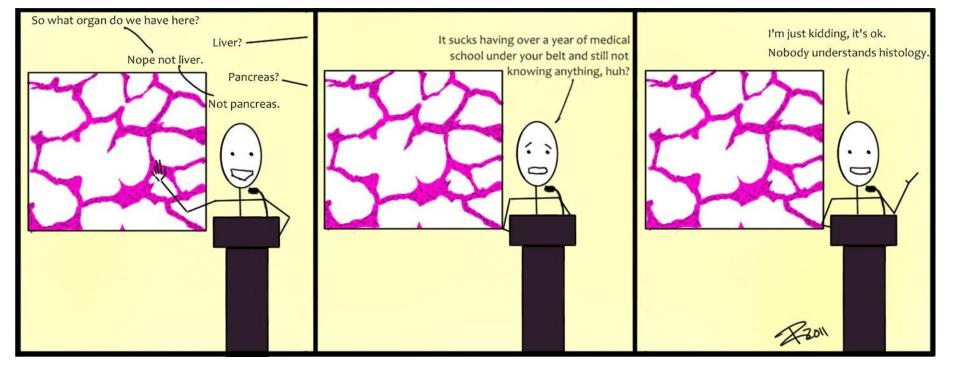
# Guide to General Histology and Microscopic Anatomy

Petr Vaňhara, Miroslava Sedláčková, Irena Lauschová, Svatopluk Čech, Aleš Hampl



Masaryk University, Brno 2017

http://www.med.muni.cz/histology



# Thank you for attention

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