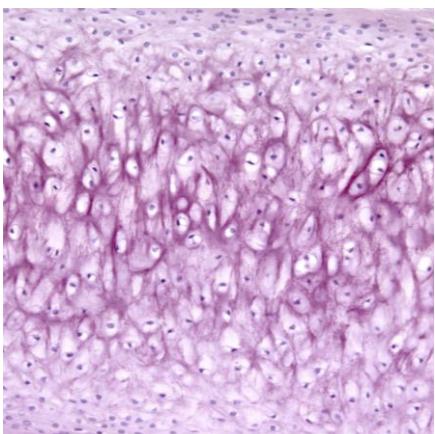
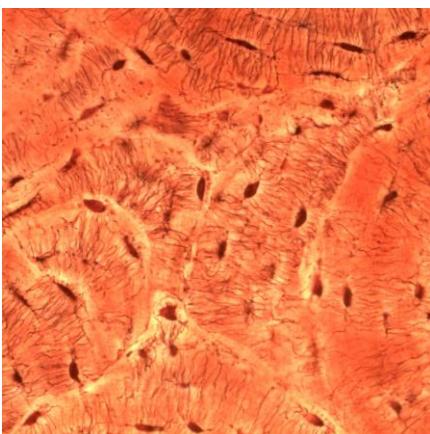
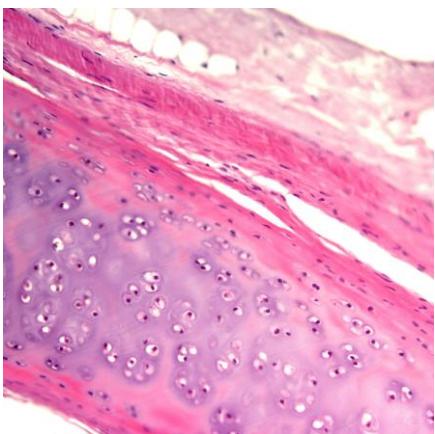


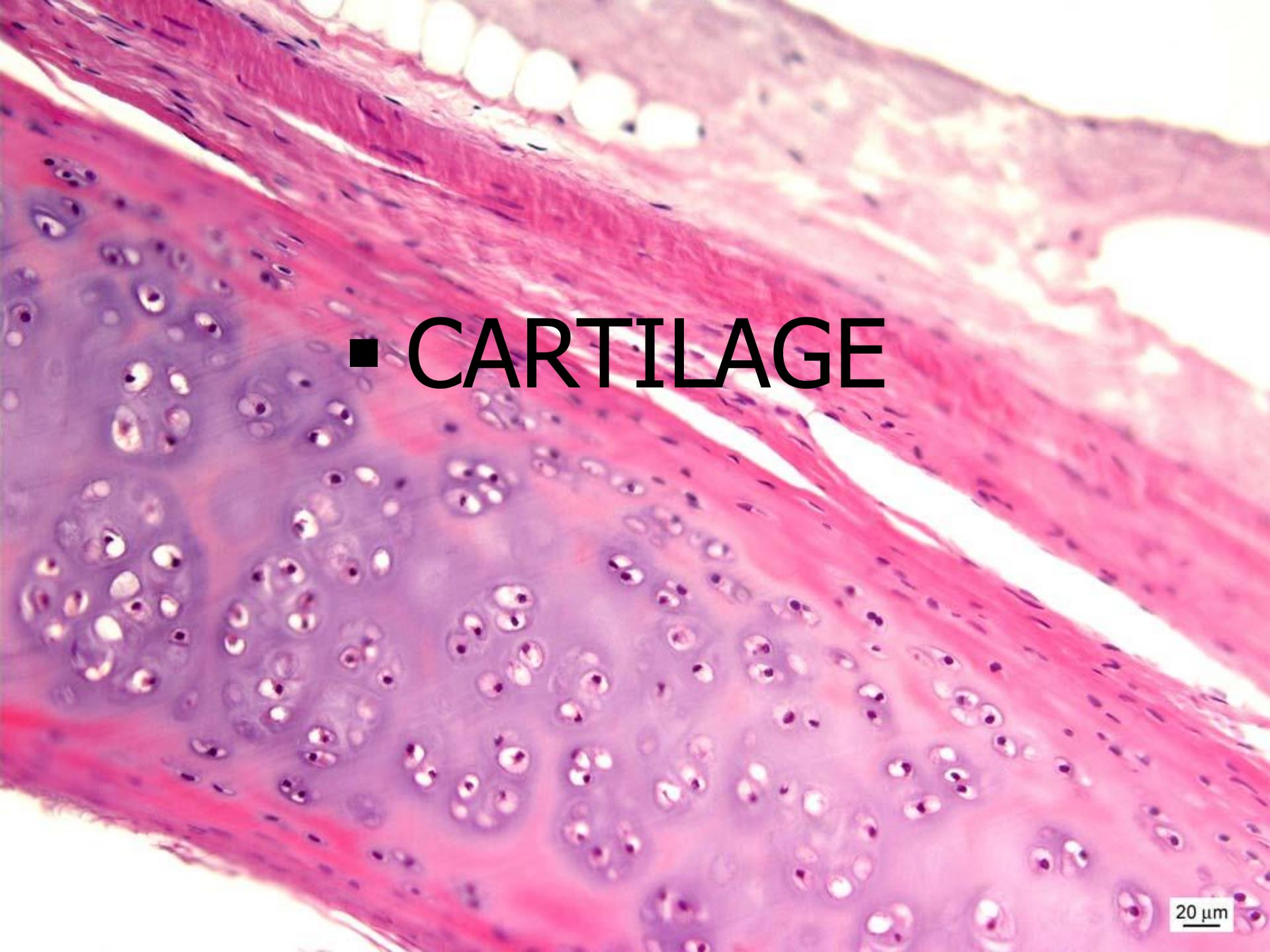
CARTILAGE AND BONE

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and Embryology,
Faculty of Medicine MU

pvanhara@med.muni.cz



A light micrograph showing a cross-section of cartilage tissue. The lower portion of the image displays a dense arrangement of chondrocytes within lacunae, stained purple. Above this, a layer of pink-stained collagen fibers runs diagonally. The top edge shows a white, granular material, likely articular cartilage. A scale bar in the bottom right corner indicates 20 micrometers.

■ CARTILAGE

20 μm

■ Cartilage

General features:

- **specialized connective tissue** with continuous ECM
- flexible, mechanically resistant
- avascular, non-innervated
- support of soft tissues - trachea, larynx
- skeletal support - costal cartilages
- diarthrosis - joints
- bone growth

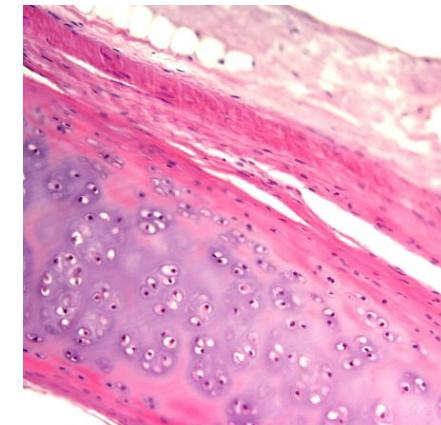


1. **cells**
2. **fibrils**
3. **amorphous ground substance**

■ Composition and structure

- Perichondrium – connective tissue around cartilage (except joints)

Nutrition
Growth



- Extracellular matrix – water, proteoglycans and collagen II fibrils

Solid consistency
Pressure elasticity

- Cells of cartilage - chondroblasts, chondrocytes

Growth
ECM production

Distribution

cartilage in adults

Hyaline

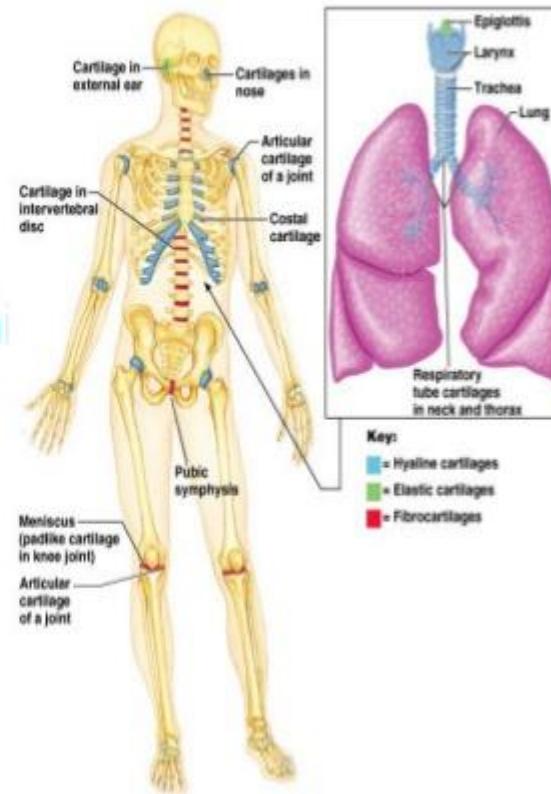
- Nose
- Joint surfaces
- Costal
- Larynx - voice box
- rings of trachea & bronchi

Elastic

- External ear
- Epiglottis
- Eustachian tube

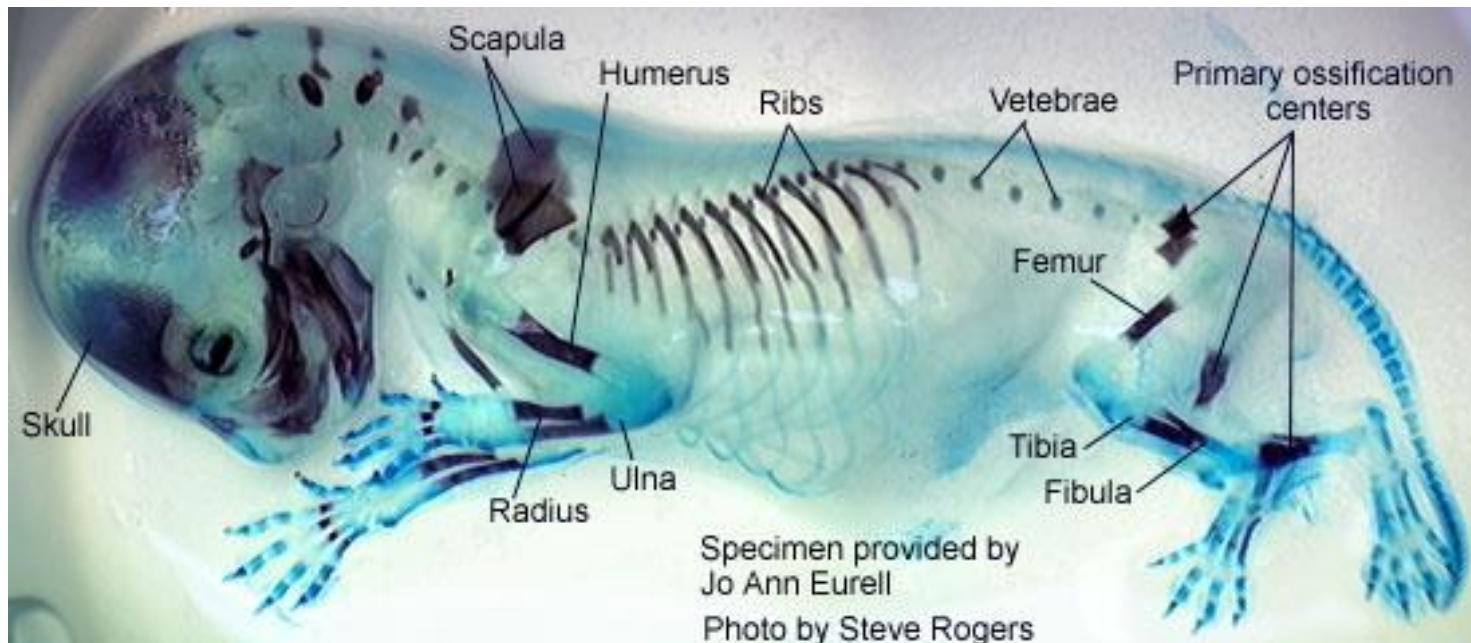
Fibrous

- IVDs
- Pubic symphysis
- meniscus in knee joint



Distribution

Hyaline



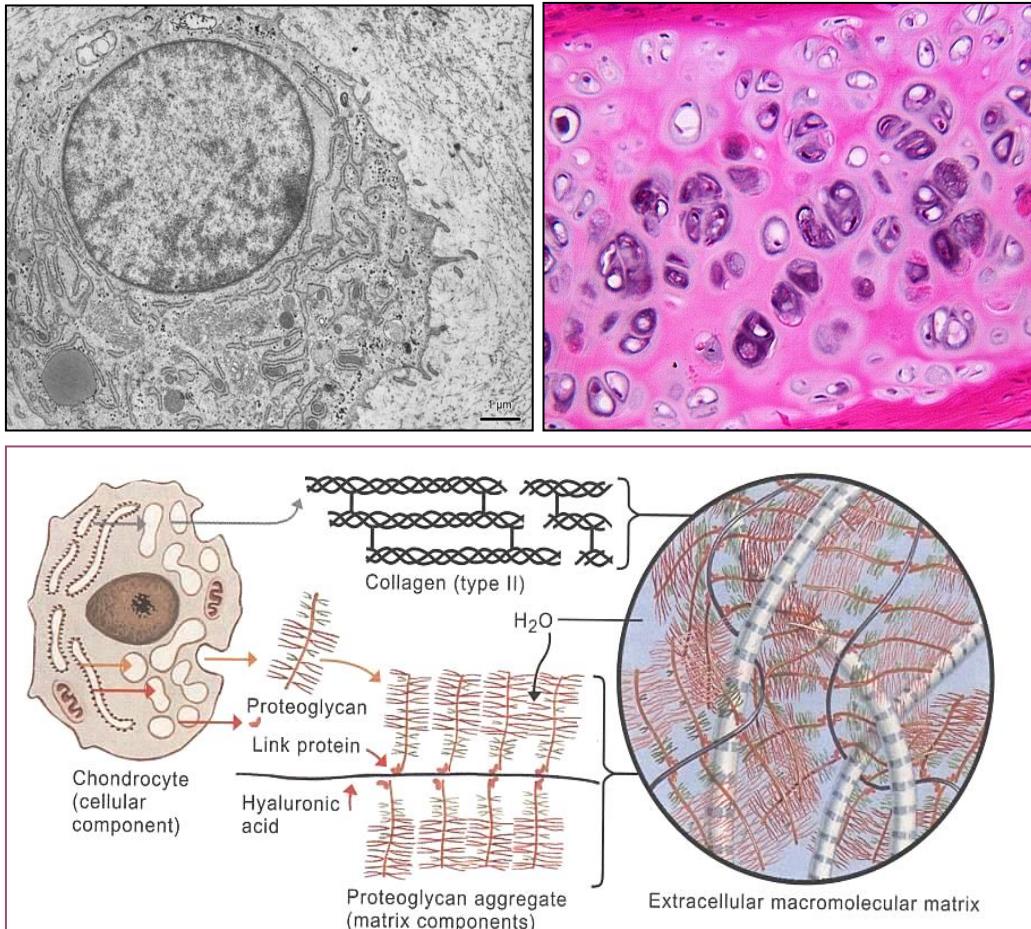
Alcian Blue&Alizarin Red

- most abundant
- temporary embryonal/fetal skeleton
- epiphyseal growth plate
- articulation (joints) respiratory passages

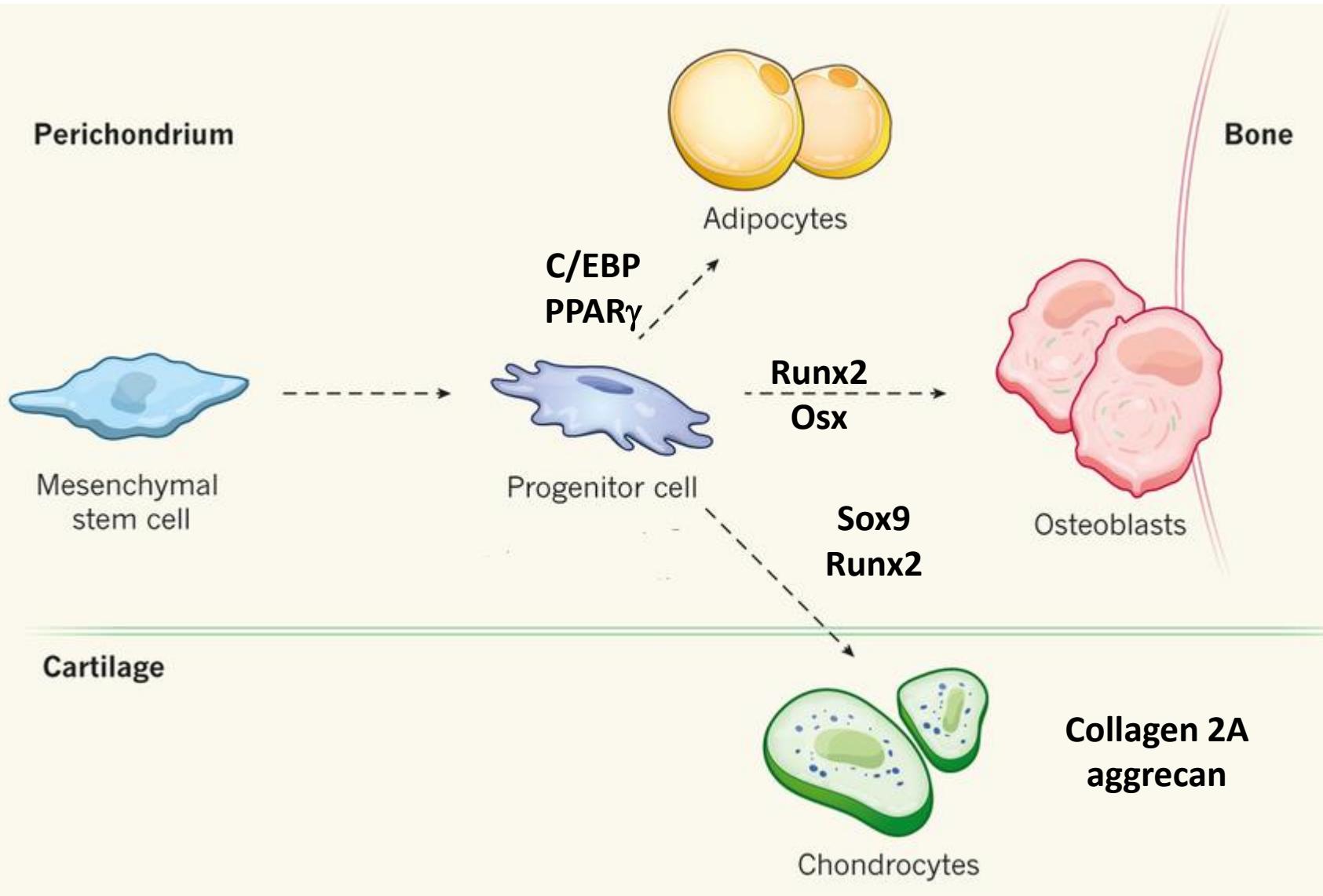
■ Cells of cartilage

■ Chondroblasts and chondrocytes

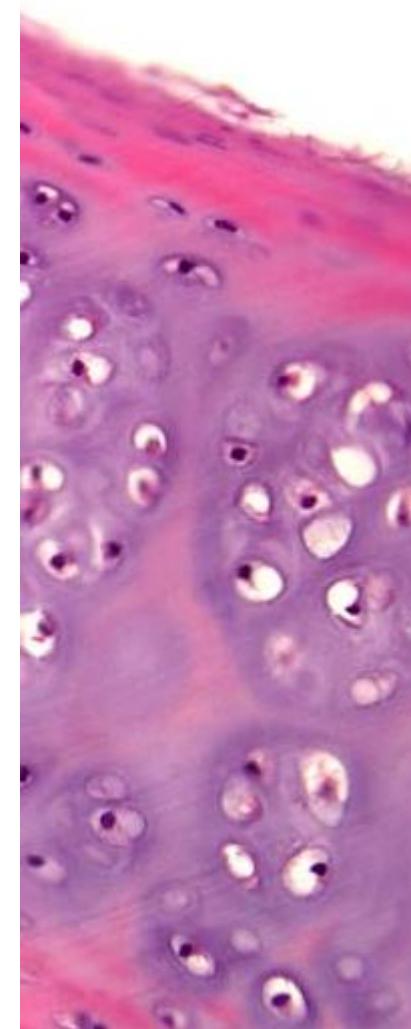
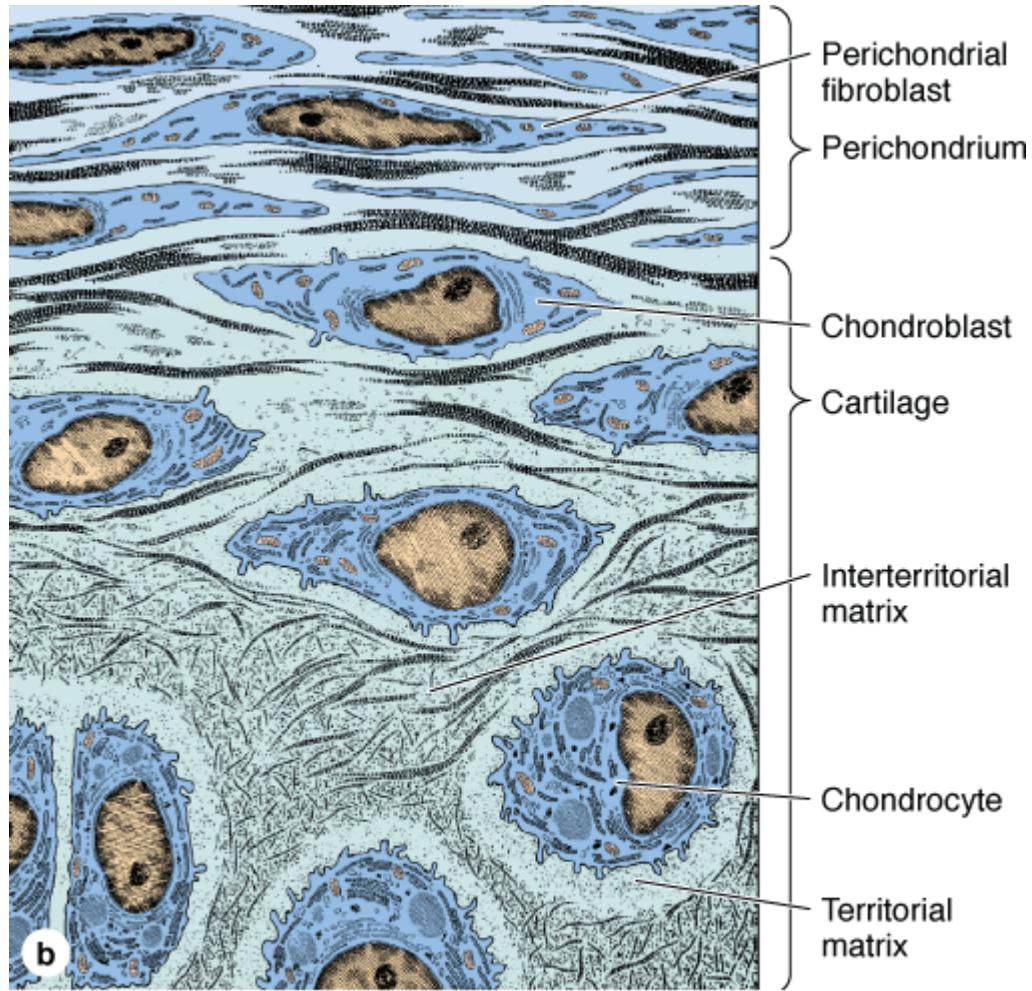
- mesenchymal origin
- typical ultrastructure of proteosynthetically active cells
- production of extracellular matrix
- interstitial proliferation
- isogenetic groups, lacunae



■ Origin of chondrocytes



■ Origin of chondrocytes

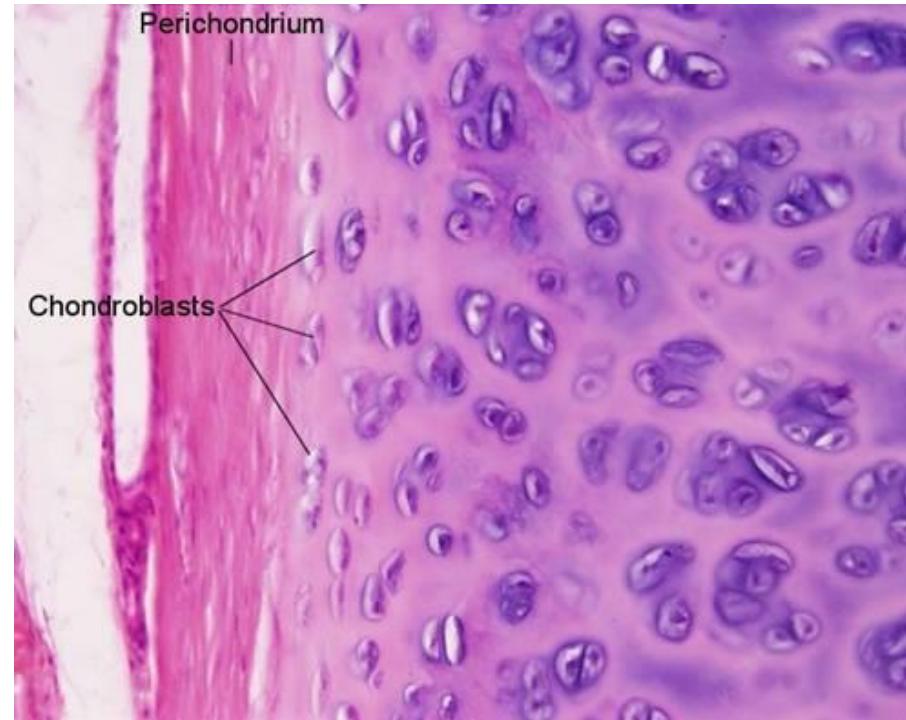


Source: Mescher AL: Junqueira's Basic Histology: Text and Atlas, 12th Edition: <http://www.accessmedicine.com>

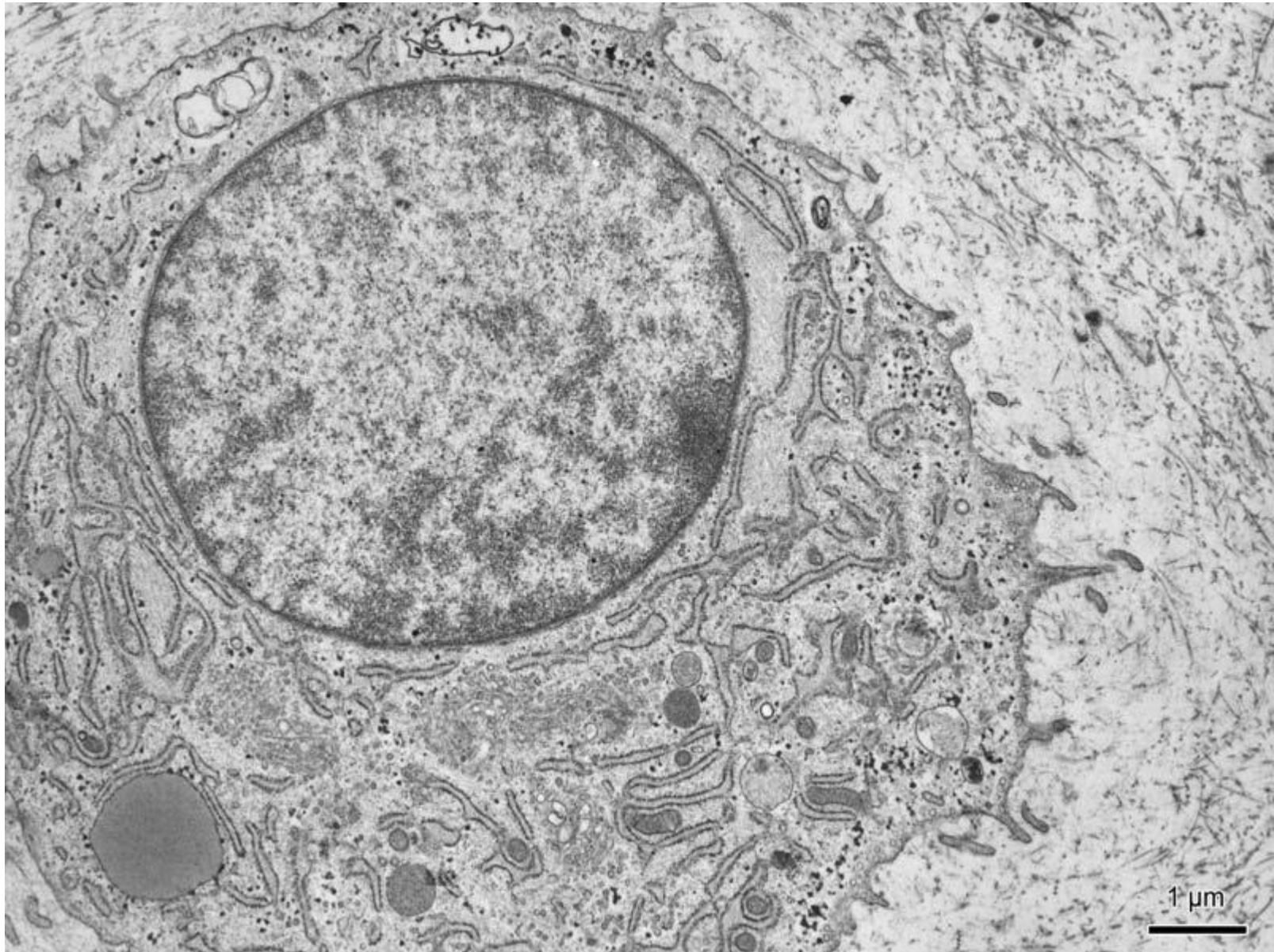
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■ Ultrastructure of chondrocytes

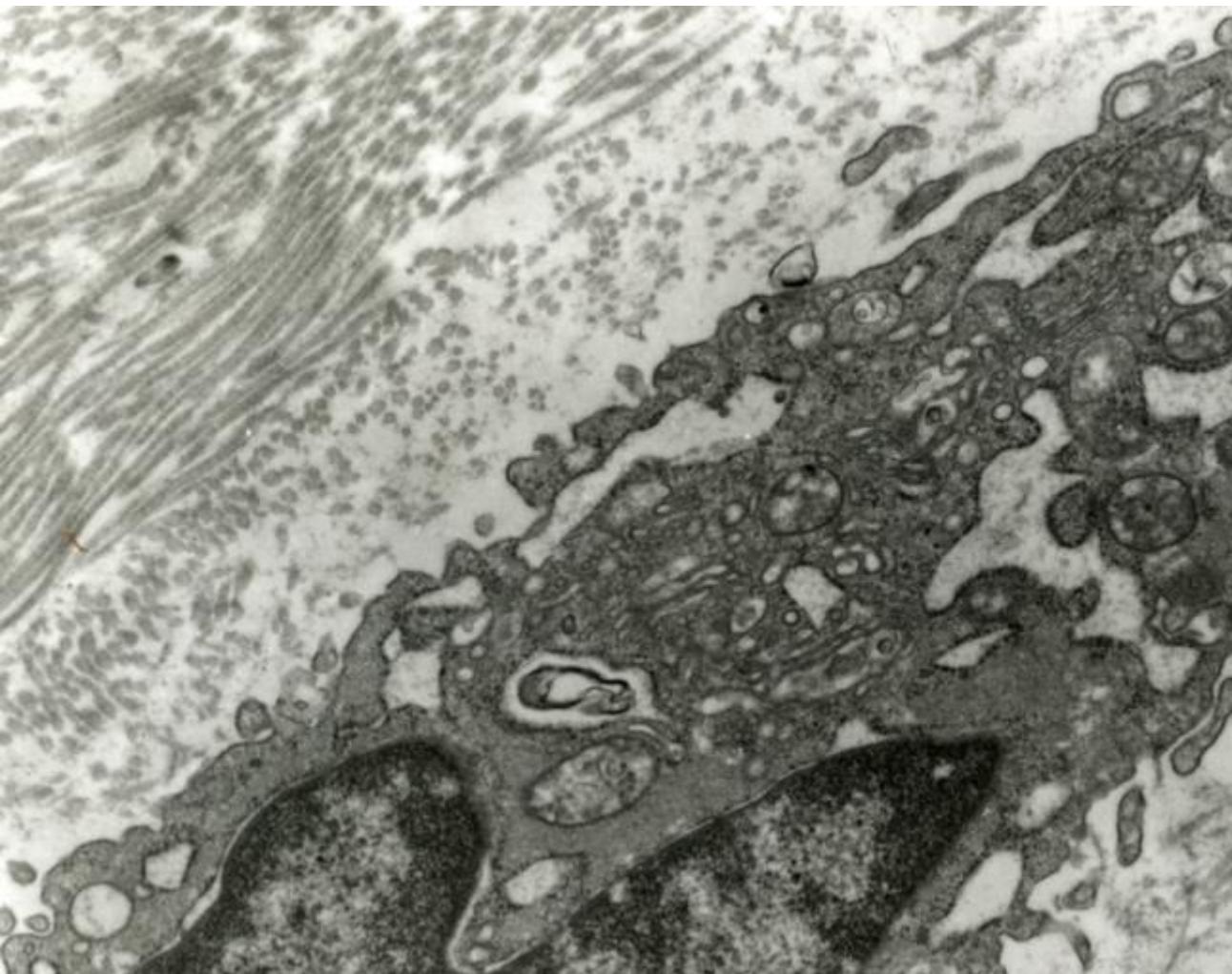
- oval → round cells
- rich in organelles, especially rER and GA
- glycogen granules (anaerobic metabolism)
- occasionally lipid droplets



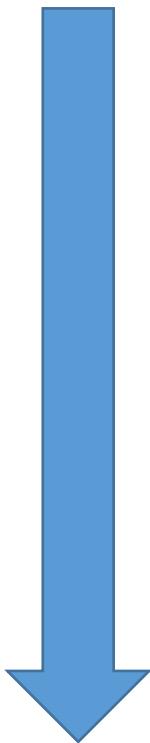
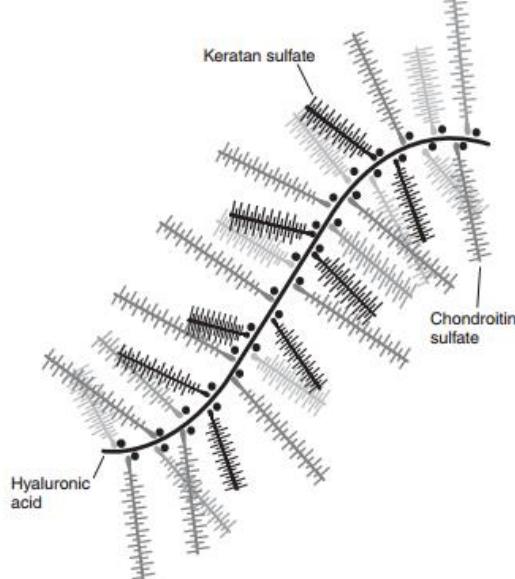
- Ultrastructure of chondrocytes



- Ultrastructure of chondrocytes



■ Extracellular matrix



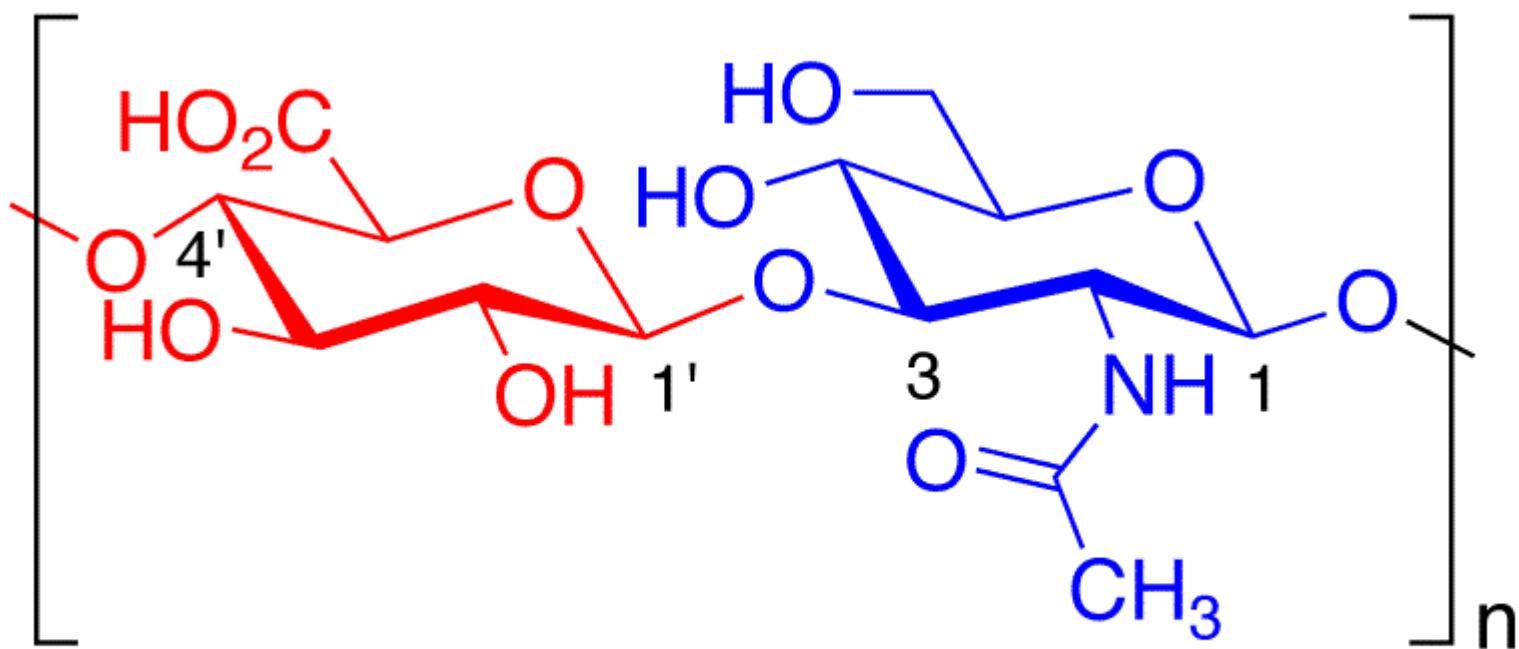
1. glycosaminoglycans
2. proteoglycans
3. fibers
4. water

biomechanical properties

Glycosaminoglycans

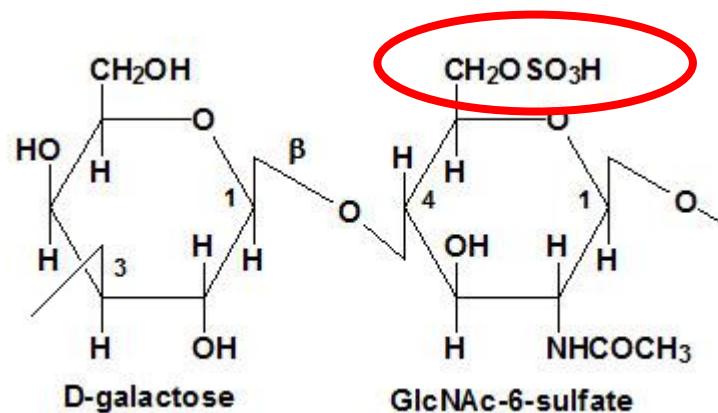
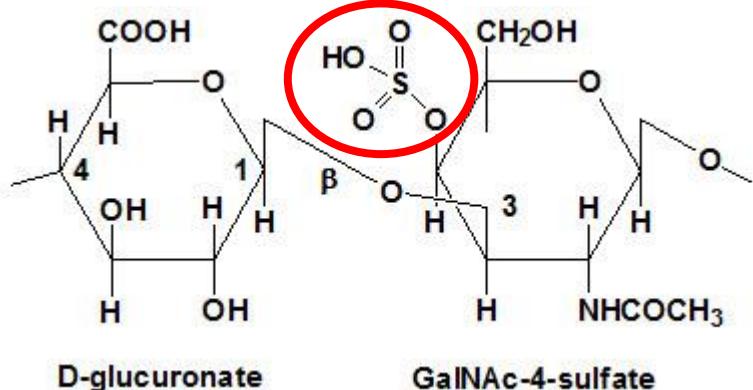
linear unbranched polysaccharides containing a repeating disaccharide unit:

1. N -acetylgalactosamine (GalNAc) or N -acetylglucosamine (GlcNAc)
2. uronic acid (glucuronate (GlcA)) or iduronate.



Glycosaminoglycan Localization

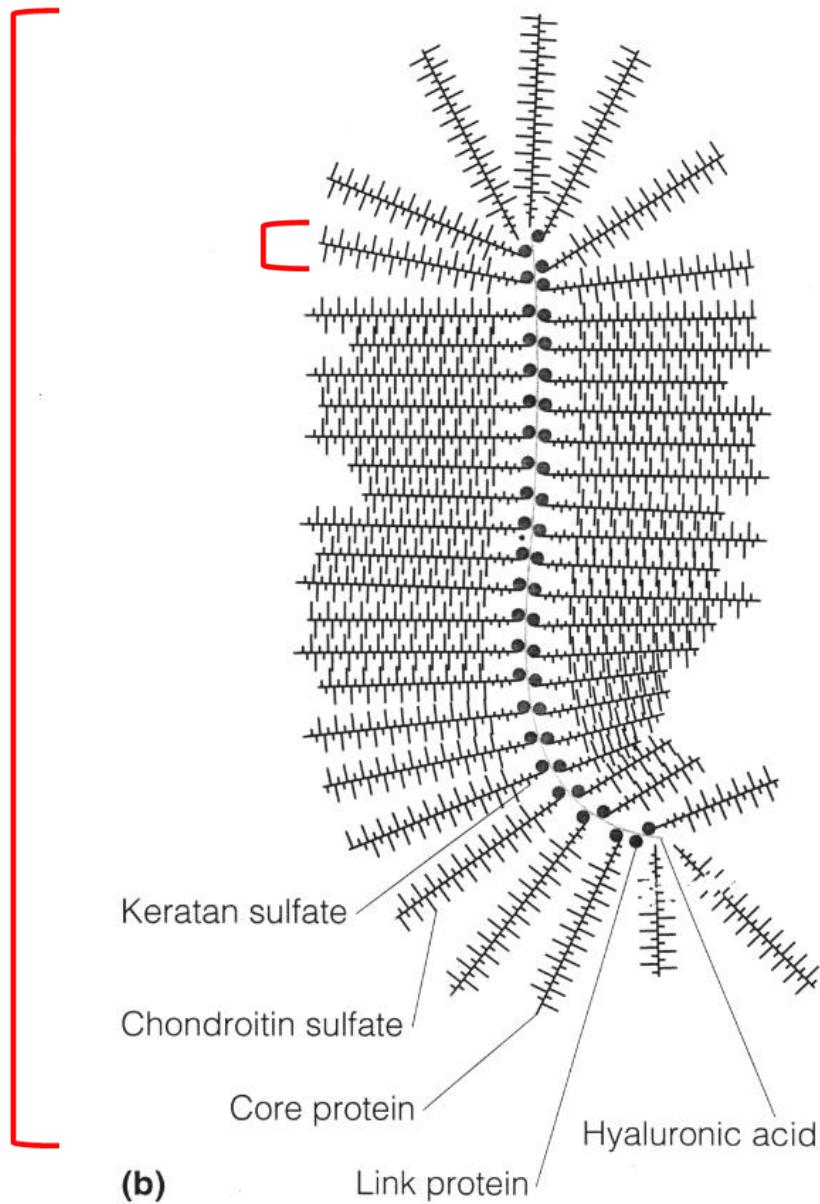
Hyaluronic acid	Umbilical cord, synovial fluid, fluid of corpus vitreum, cartilage
Chondroitinsulphate	Cartilage, bone, cornea, skin, notochord, aorta
Dermatansulphate	Skin, ligaments, adventitia of aorta
Heparansulphate	Aorta, lungs, liver, basal membranes
Keratansulphate	Iris, cartilage, nucleus pulposus, anulus fibrosus



Proteoglycans

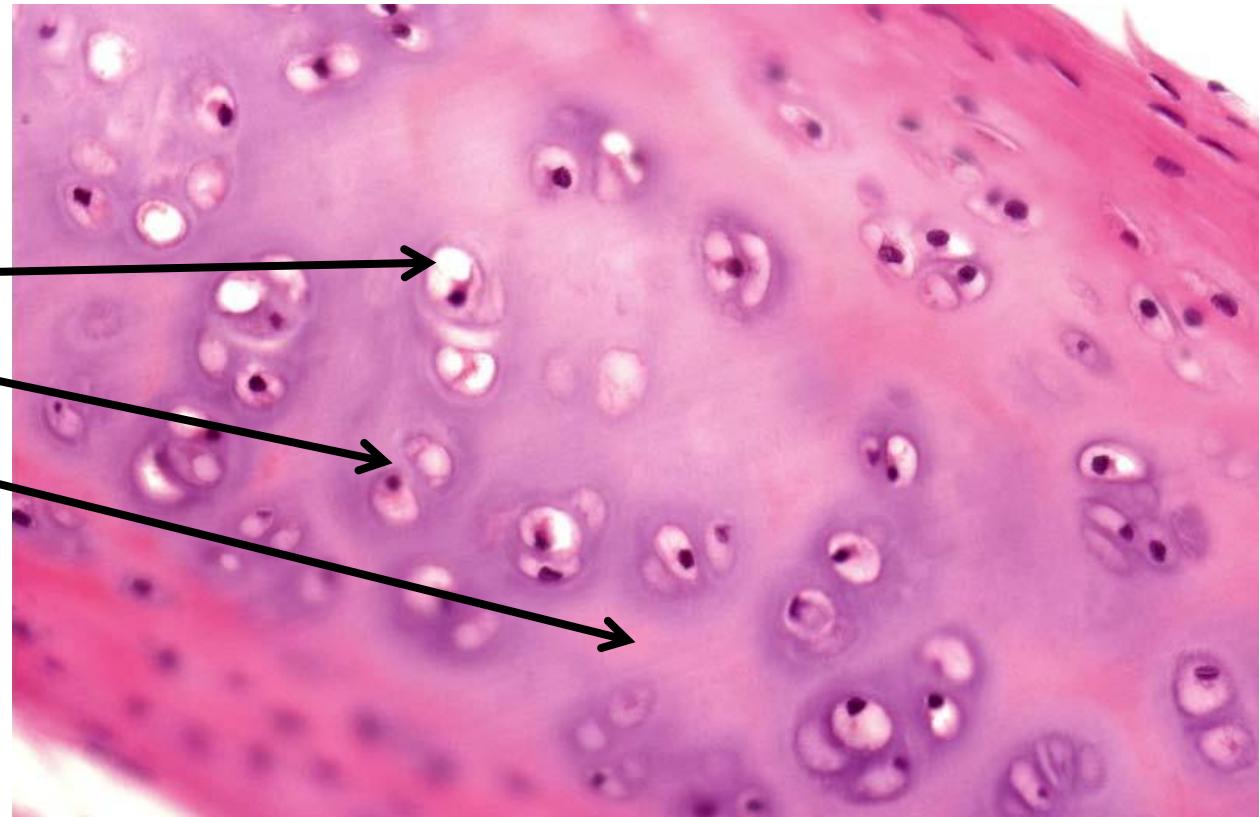
- protein + dominant linear saccharide component
- proteoglycan aggregates
- water-binding, volume dependent of hydration
- **aggrecan (cartilage)**
- syndekan
- fibroglykan

Figure 9.25b Proteoglycan structure in bovine cartilage



■ Architecture of extracellular matrix

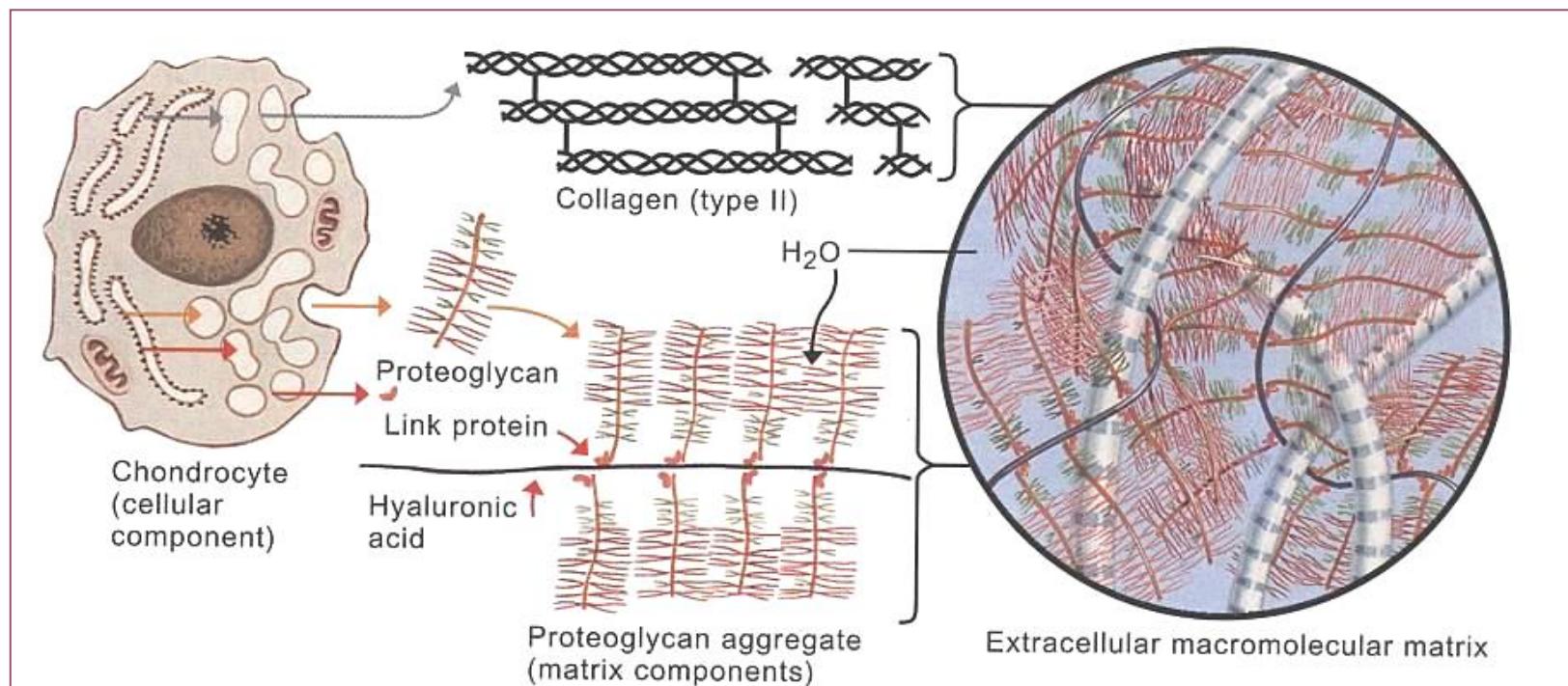
- pericellular
- territorial
- interterritorial



transduction of biochemical and biomechanical signals

■ Extracellular matrix

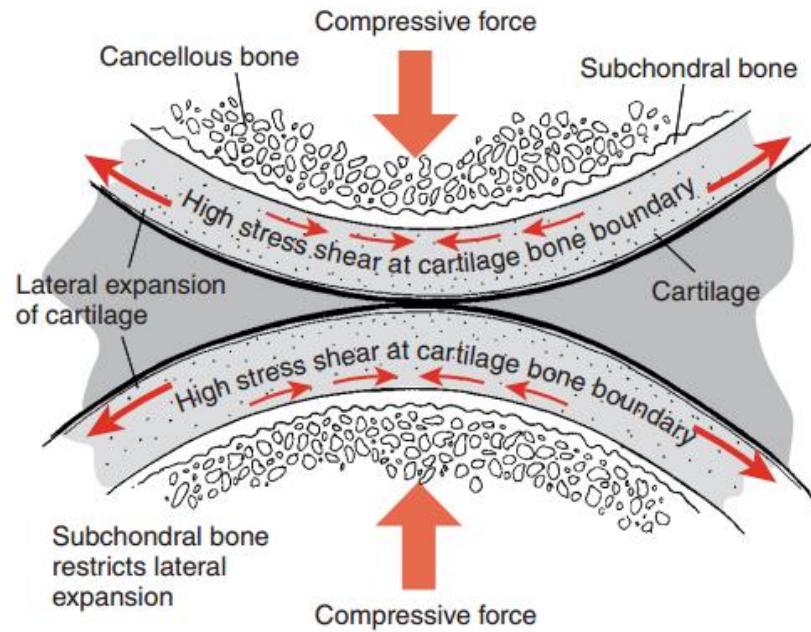
- **collagen fibrils**
- col II + col IX/XI
- thin fibrils (15-20 nm → no striation) do not form fibers
- interconnected with perichondrium
- **proteoglycans and glycosaminoglycans**
- aggrecan hyaluronan-based aggregates
- water
- 80%



■ Architecture of extracellular matrix

- **pressure elasticity**

- proteoglycans – polyanionic (COO^- , SO_4^{2-})
- expansion prevented by collagen fibrils
- repulsion forces



- **biphasic model of cartilage conditioned by ECM composition**

- proteoglycans, collagen, cells, and lipids constitute the solid phase of the mixture
- interstitial fluid that is free to move through the matrix (fluid phase)
- under impact loads, fluid flows through the framework, until the cartilage starts to behave as a single-phase, incompressible, elastic solid - the fluid does not flow
- after load release, fluid returns
- nutritive aspect

■ Architecture of extracellular matrix

- **synovial cartilage**

I. tangential (superficial) zone

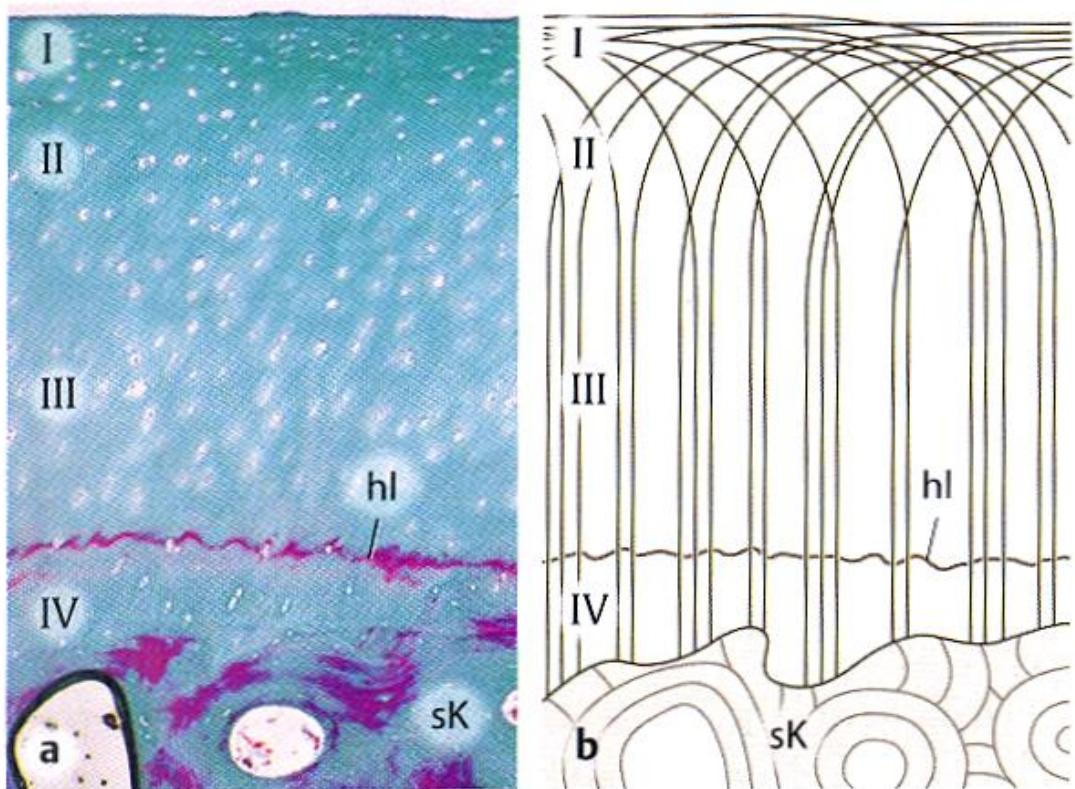
II. transitional zone

III. radial (deep) zone

tide mark

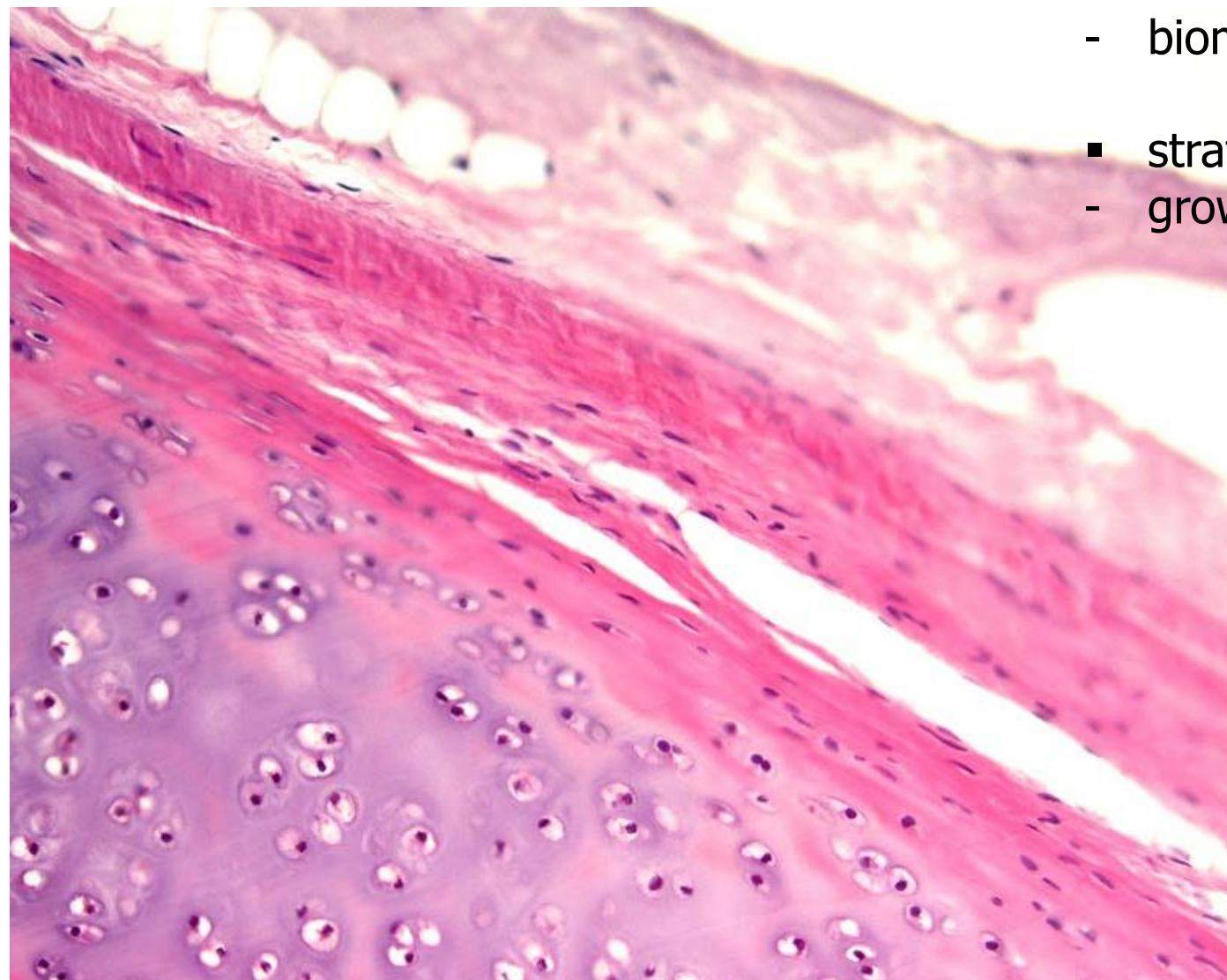
I. mineralized cartilage zone

subchondral bone



■ Perichondrium

- stratum fibrosum
 - biomechanics
- stratum chondrogenicum
 - growth



Hyaline cartilage, trachea

Perichondrium

Chondroblasts

Exchange of metabolites

Apositional growth

Isogenous (nest) cells

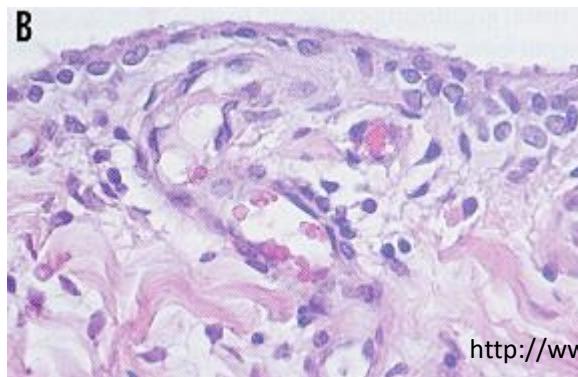
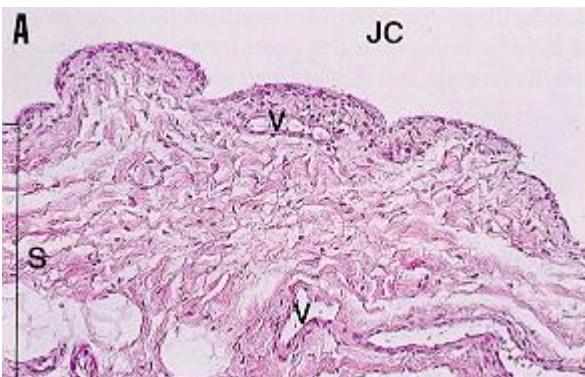
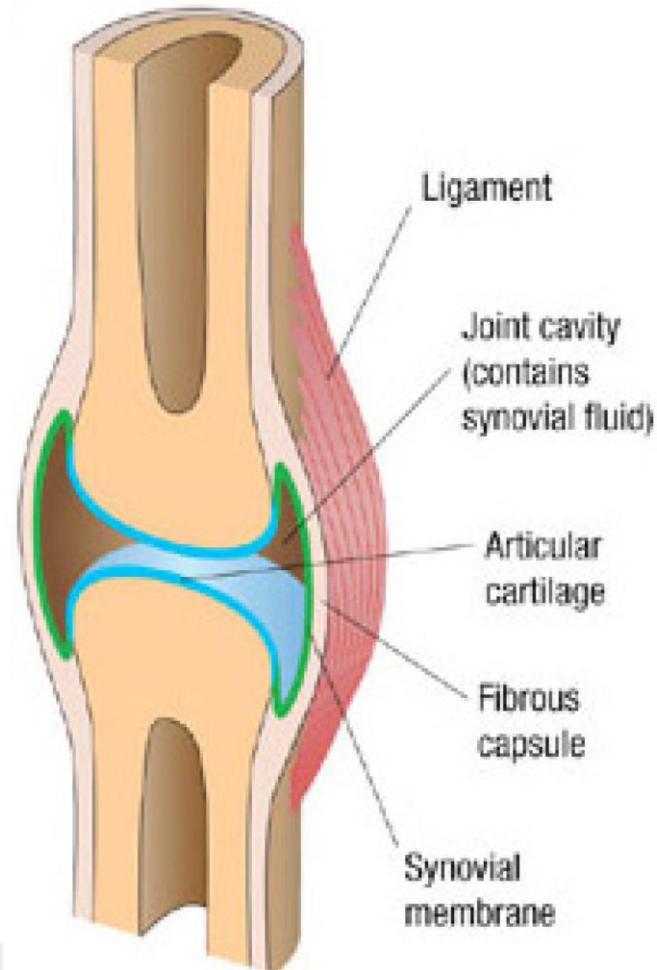
Interstitial proliferation



100 μm

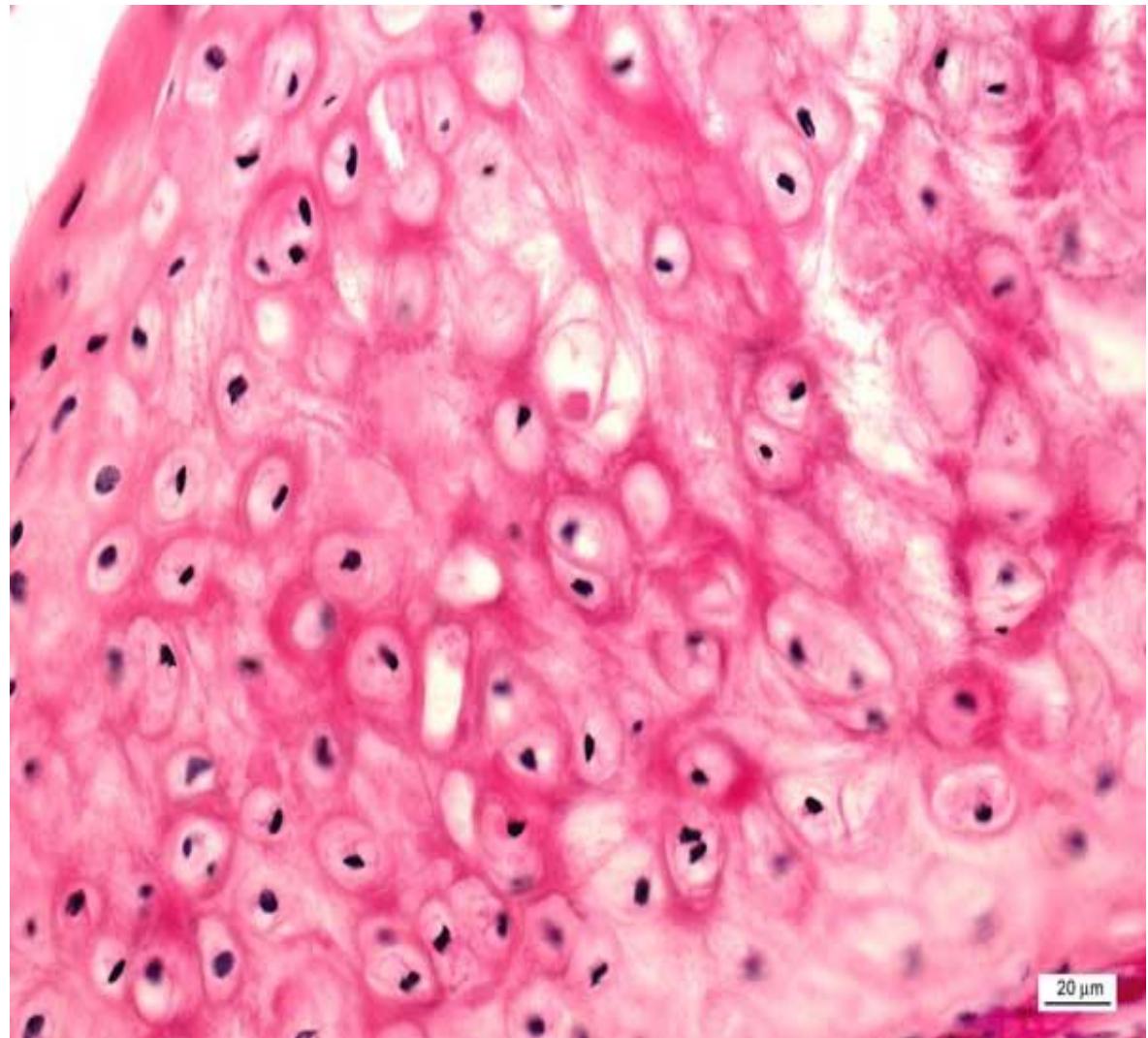
■ Synovium

- *membrana fibrosa*
 - dense collagen c.t.
- *membrana synovialis*
- intima, subintima
 - folds extending to the joint cavity
 - numerous blood and lymphatic vessels, nerves
 - discontinuous cell layers (synovialocytes)
 - basal membrane and intercellular junctions absent - **not an epithelium**
 - but mesenchymal (c.t.) origin
 - synovial fluid rich in hyaluronans
 - *bursae synoviales, vaginae tendineum*



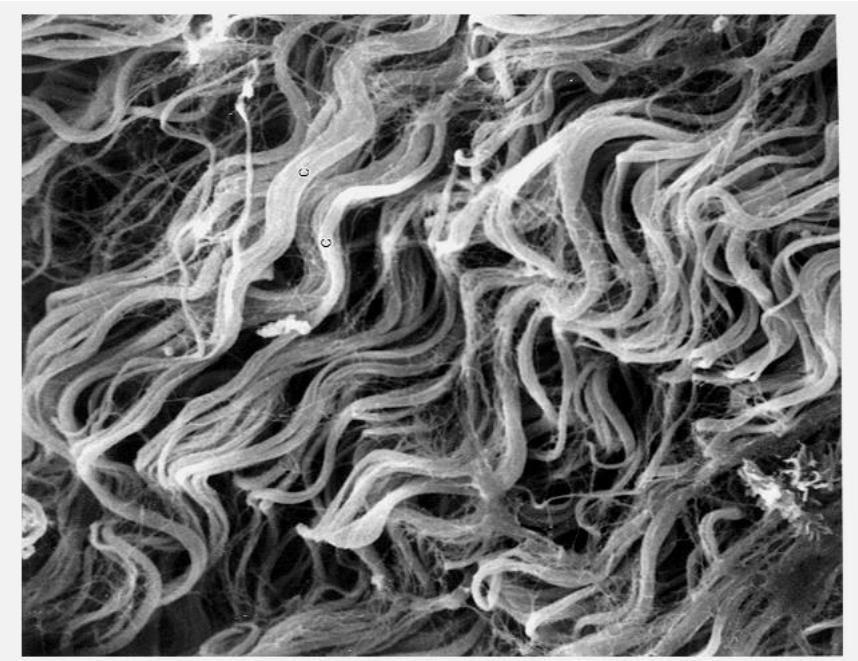
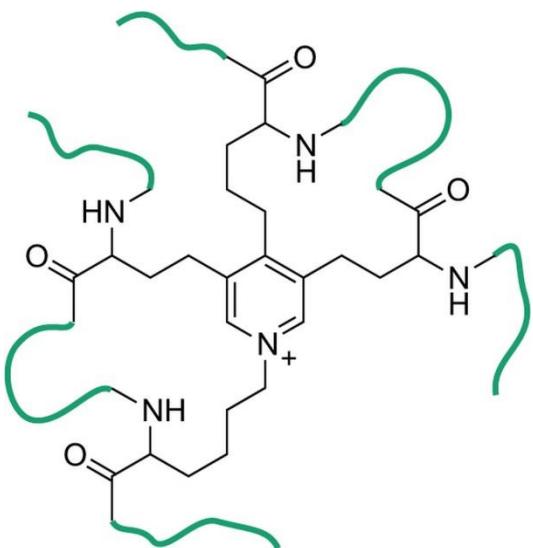
■ Elastic cartilage

- acidophilic elastic fibers in matrix
- no isogenetic groups
- auricula, meatus, larynx, epiglottis

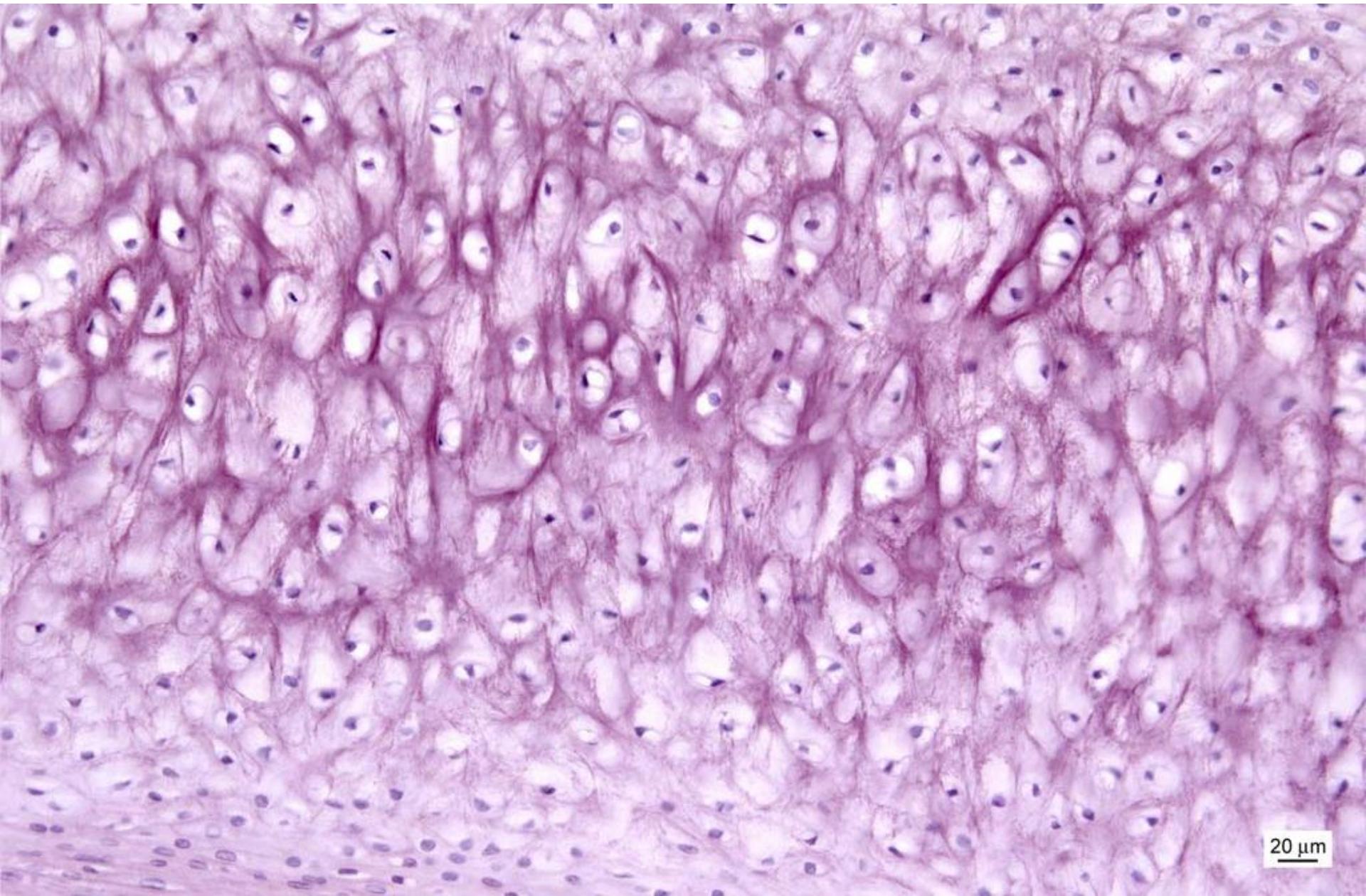


■ Elastic fibers

- 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



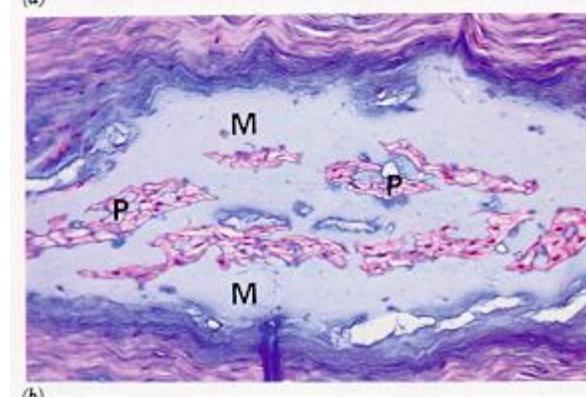
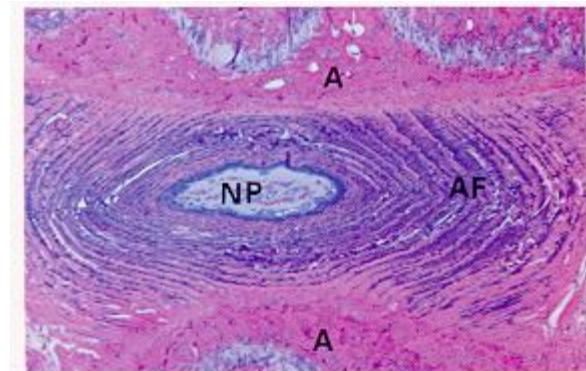
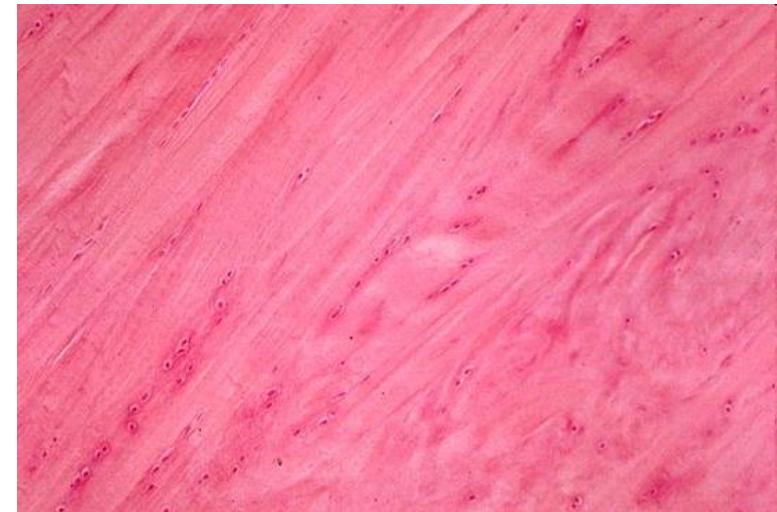
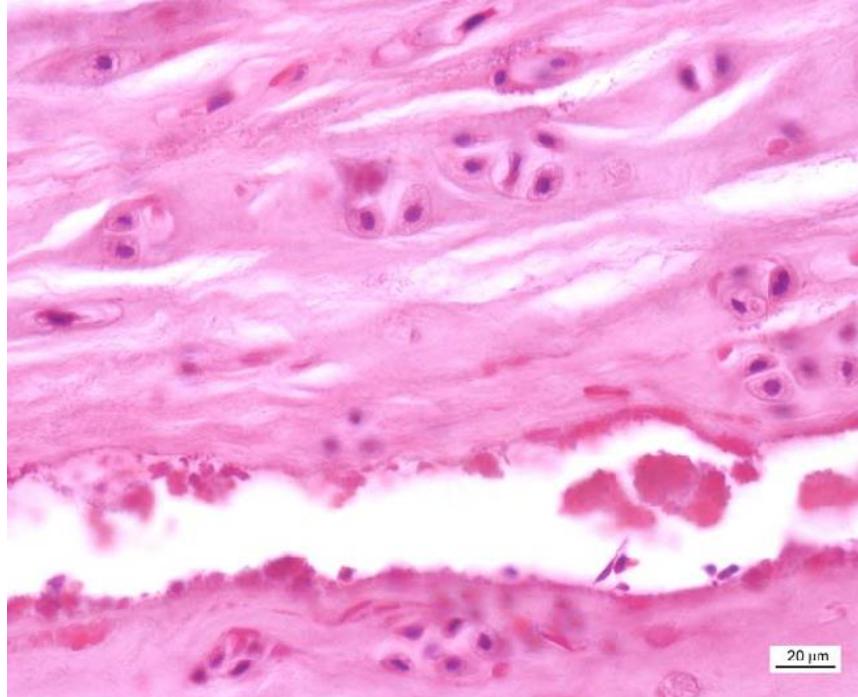
- Elastic cartilage

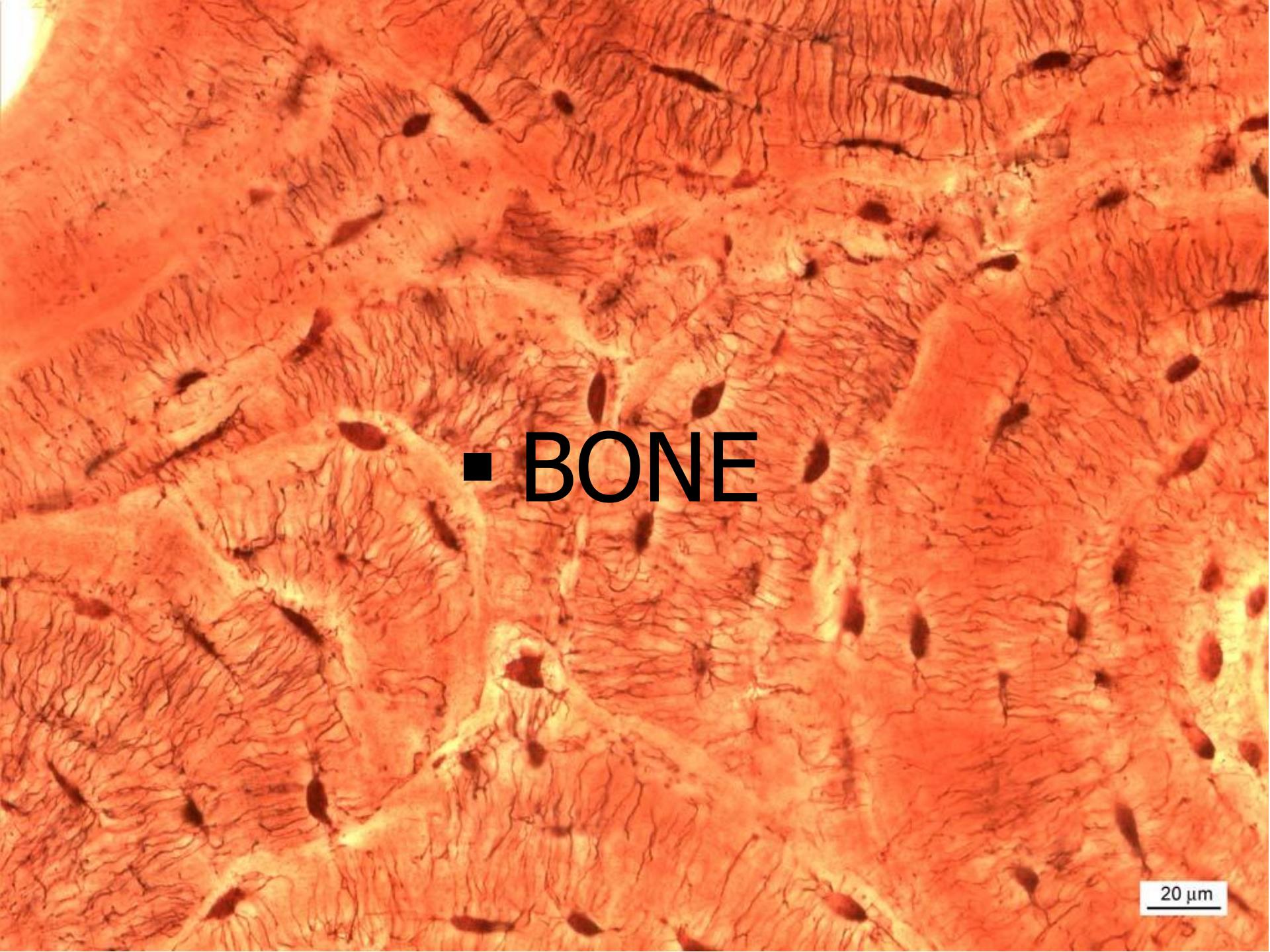


20 μm

■ Fibrocartilage

- fibrous compound dominant – collagen I and II – mechanical durability
- minimum of amorphous matrix-fibers visible
- intervertebral discs, symphysis pubis, articular discs, meniscus





A light micrograph showing a dense network of red-stained, wavy fibers, characteristic of collagenous tissue. Scattered throughout are numerous small, dark, circular structures, likely osteocytes or cellular debris. A prominent, larger, irregularly shaped area of similar staining is labeled "BONE".

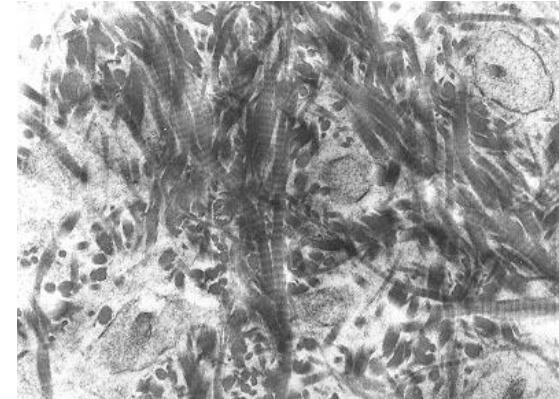
■ BONE

20 μm

■ Histological classification of bone tissue

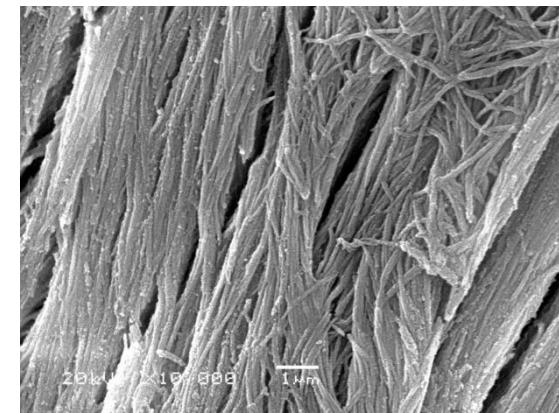
- **Primary (woven, fibrous)**

- Temporary, growth and regeneration of bones, collagen fibrils woven
- Replaced by secondary bone
- Remains only in some parts of body - sutures of skull, *tuber osseum*, tooth cement



- **Secondary (lamellar)**

- Lamellae – collagen fibers in concentric layers (3-7 μm) around a canal with capillaries = Haversian system (osteon)

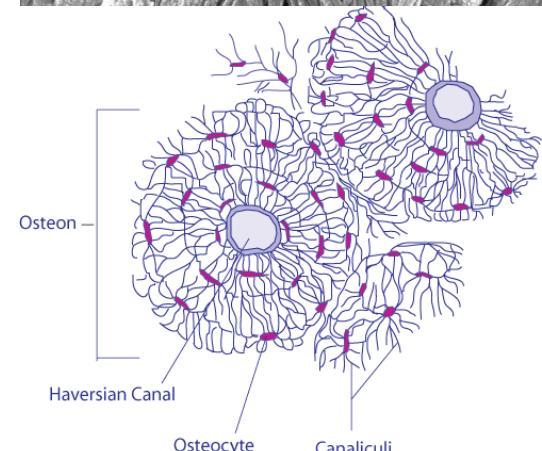


- **Spongy (trabecular)**

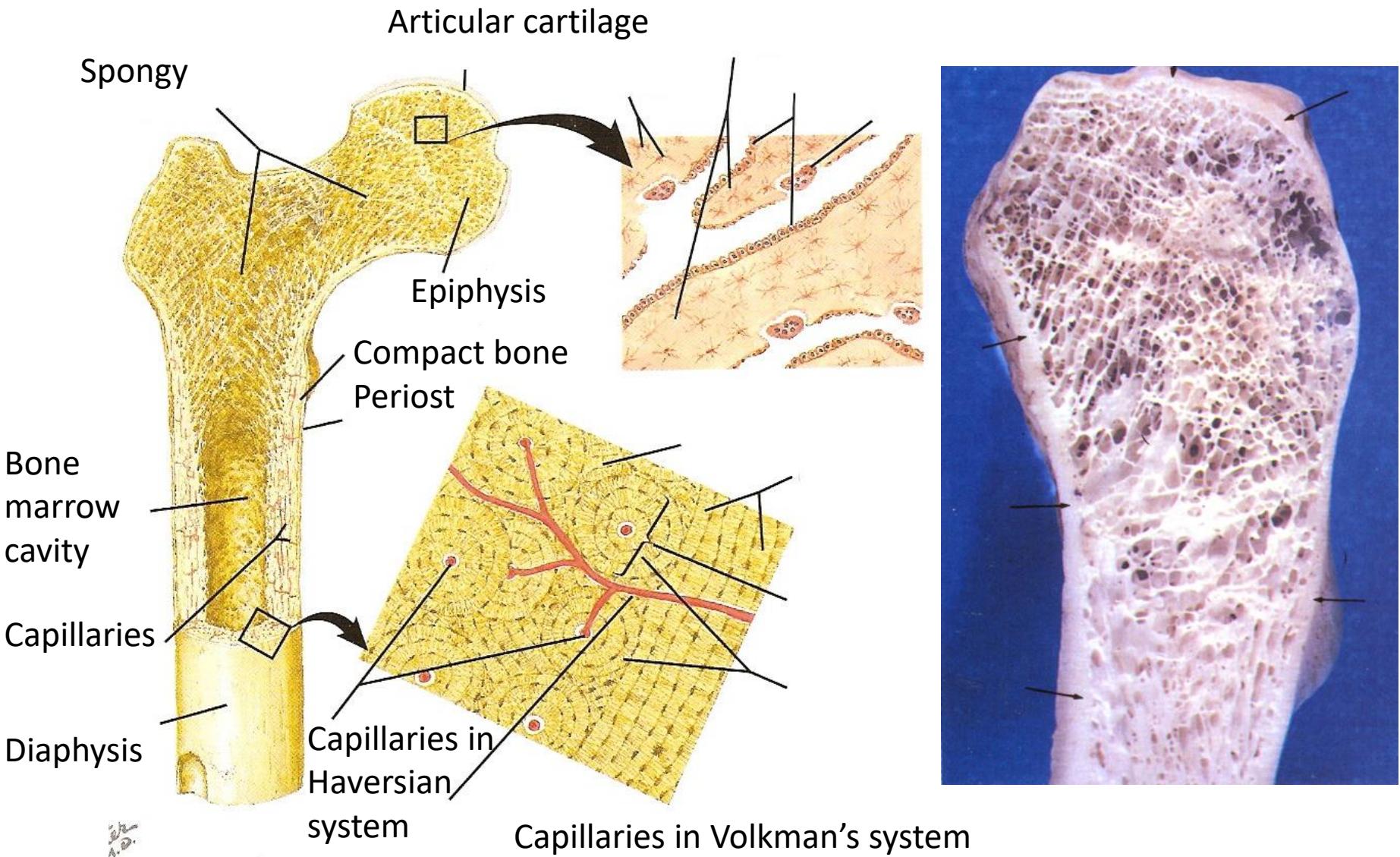
- Trabeculae, similar to compact
- Epiphyses of long bones, short bones, middle layer of flat bones of the skull (*diploe*)

- **Compact**

- Outer and inner coat lamellae typical Haversian systems
- Volkmann's canals
- Interstitial canals

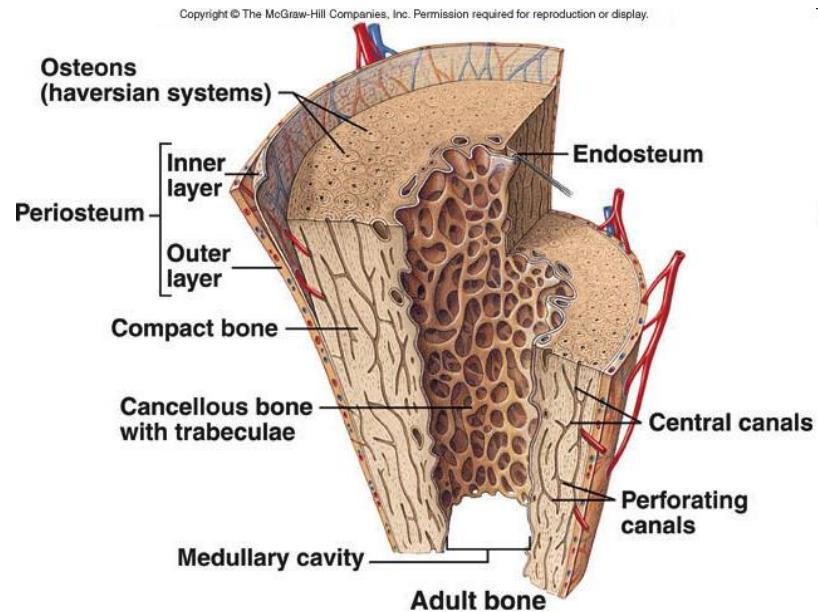


Bone

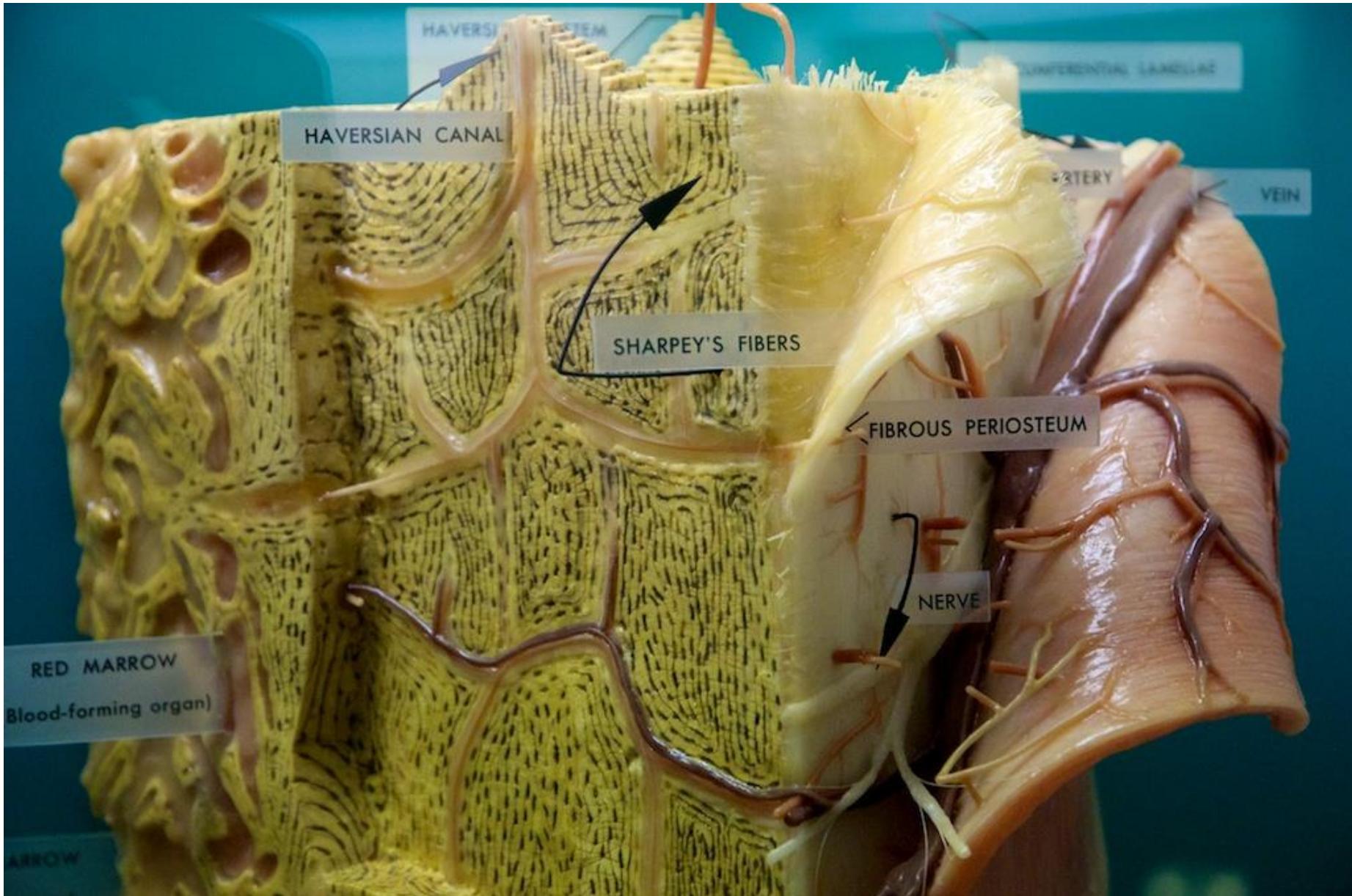


■ Surface of compact bone

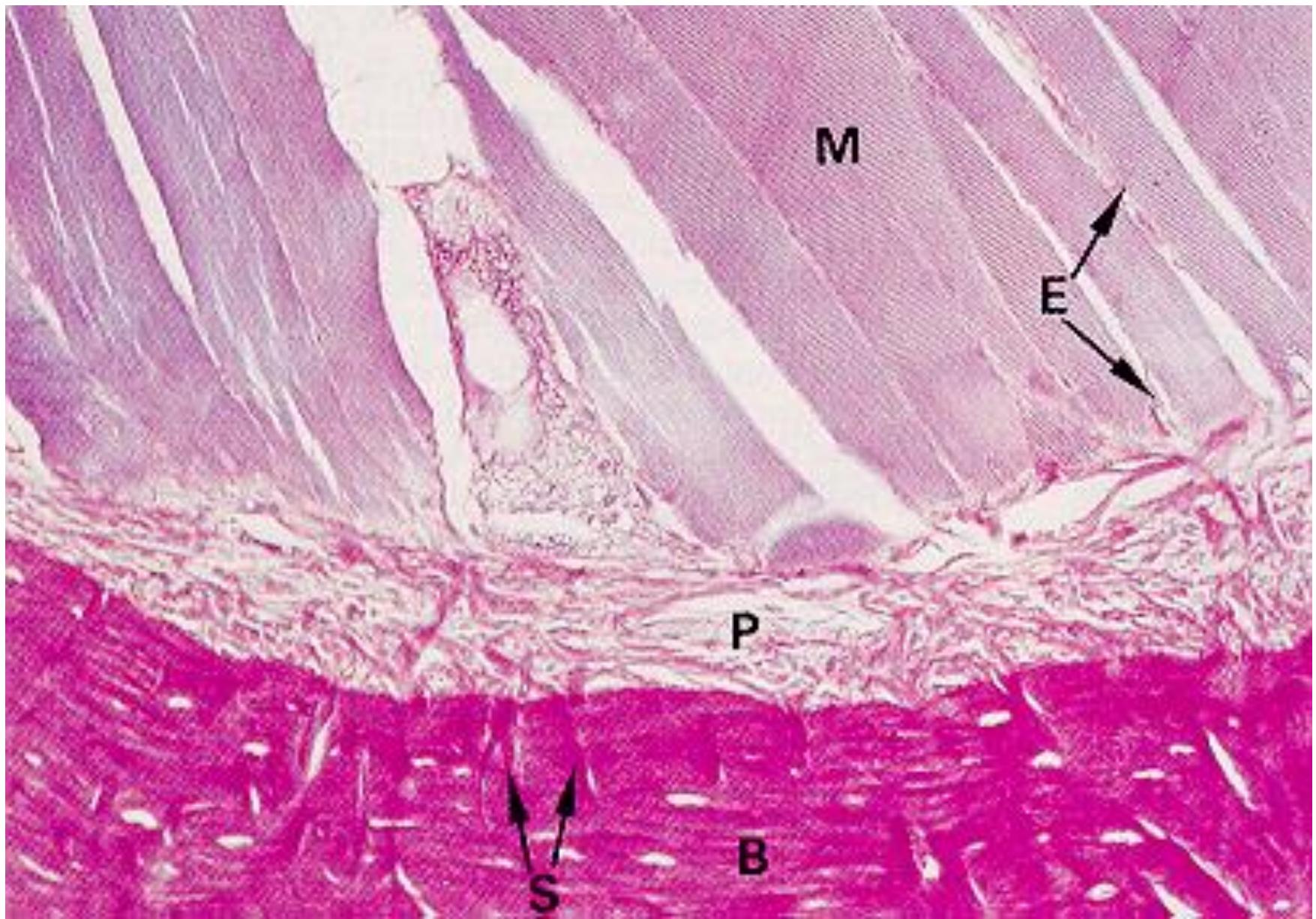
- Outer surface
- Synovial joint – hyaline cartilage
- **periosteum (periost)** – membrane – dense CT, inner layer (osteoblasts) and outer layer (fibrous CT)
- Inactive bone - fibrous CT in periost dominant
- Collagen fibers – parallel to the bone surface
- Sharpey's fibers fix periost to the bone



■ Surface of compact bone

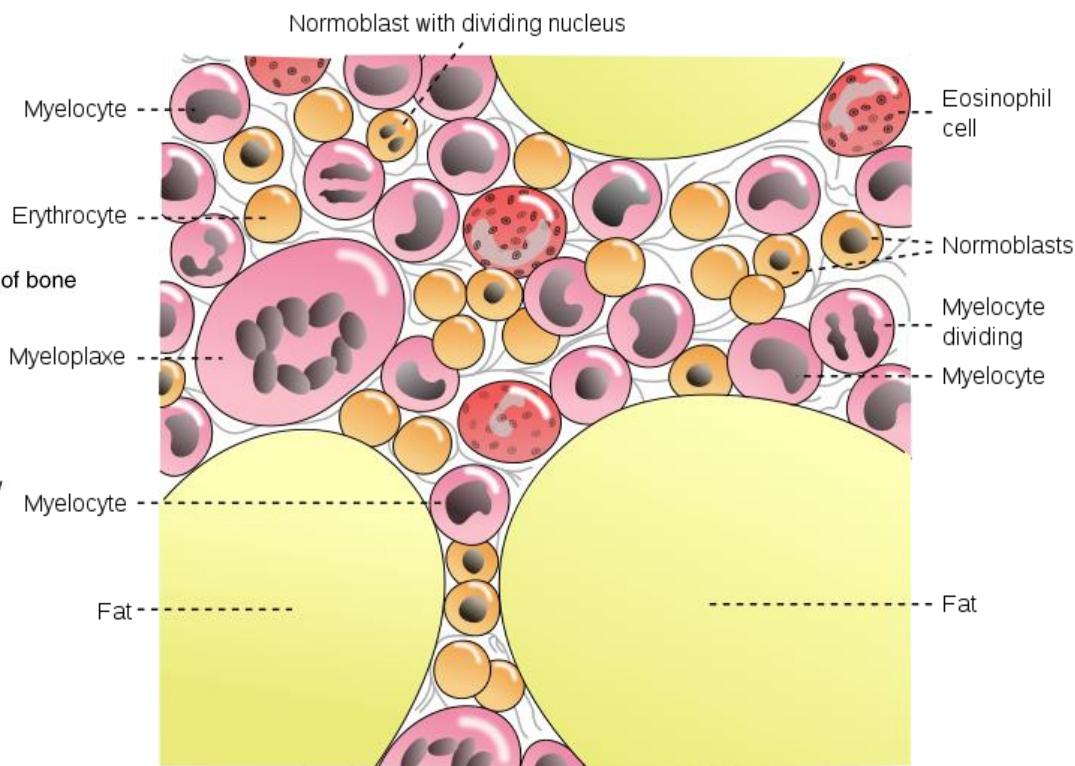
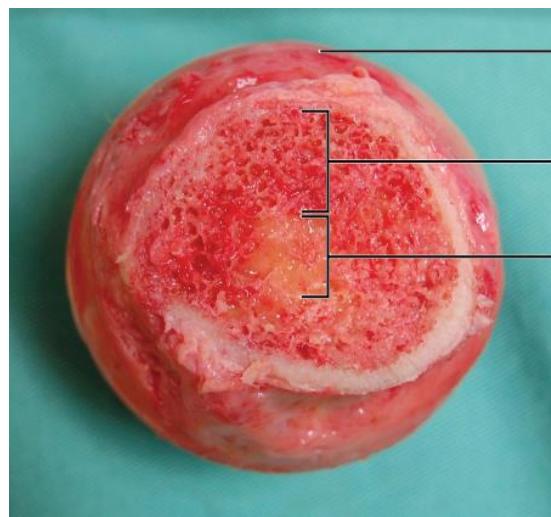
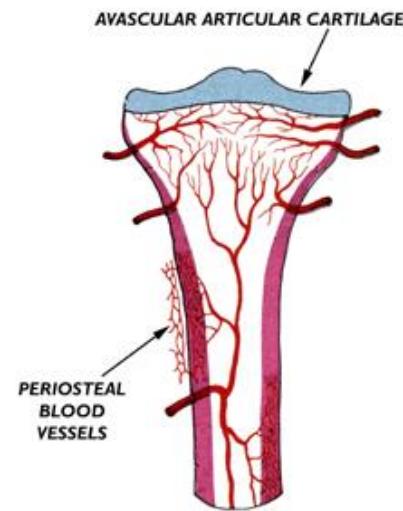


- Surface of compact bone

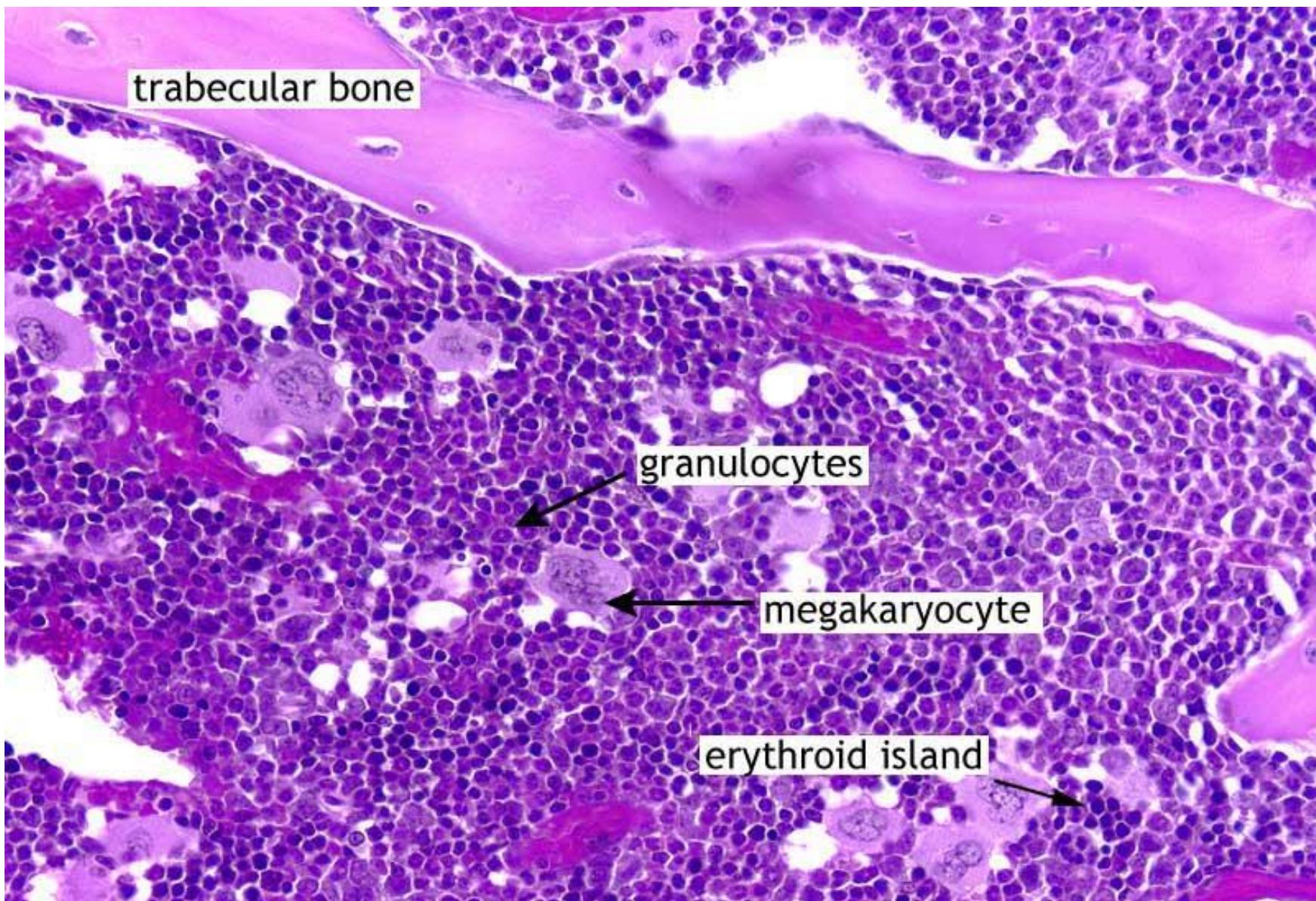


• Inner surface

- lining of medullar cavity
- endosteum (endost) – single cell lining – bone remodeling
- red bone marrow – hematopoiesis
- yellow and gray bone marrow – adipocytes or CT
- rich vascularization
- hematopoietic niche

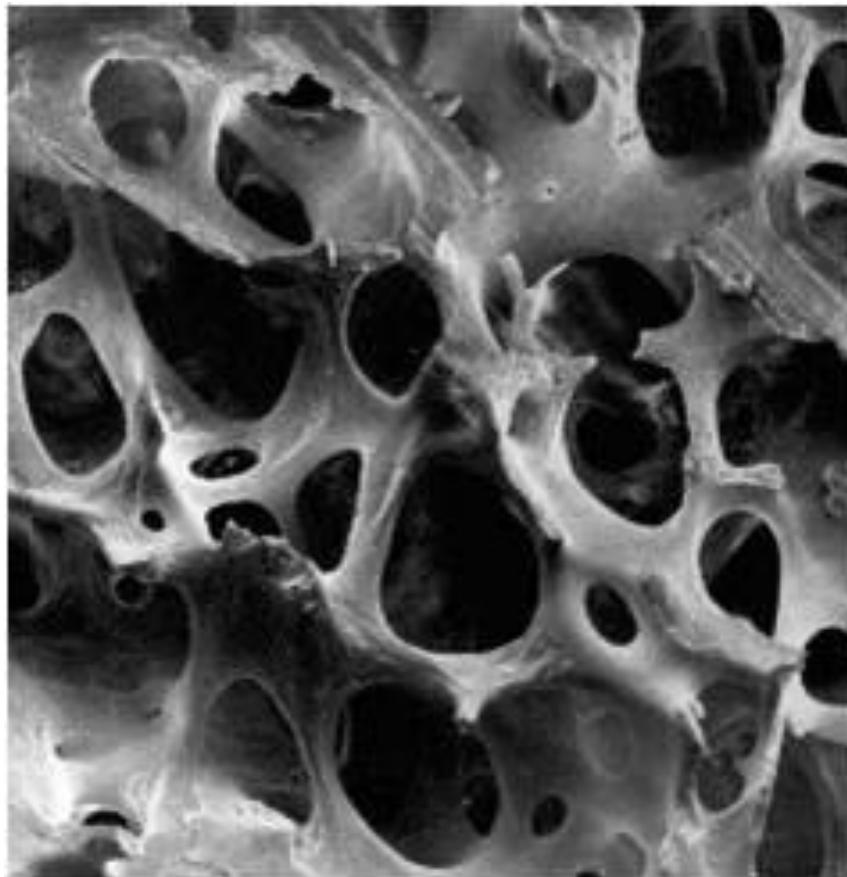


■ Endosteal surface of compact bone



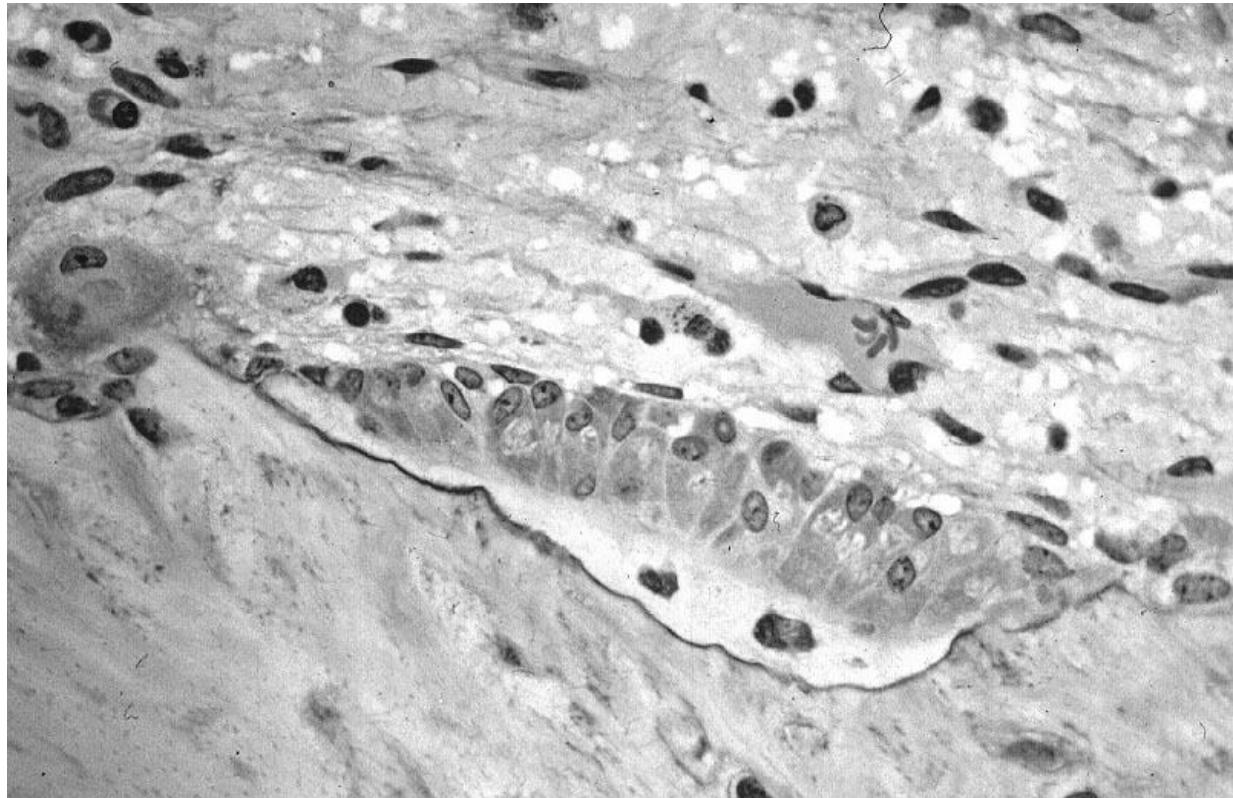
- Bone matrix

- 60% mineral compound, 24% organic compound 12% H₂O, 4% fat
- crystals – calcium phosphate, hydroxyapatite

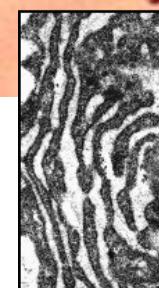
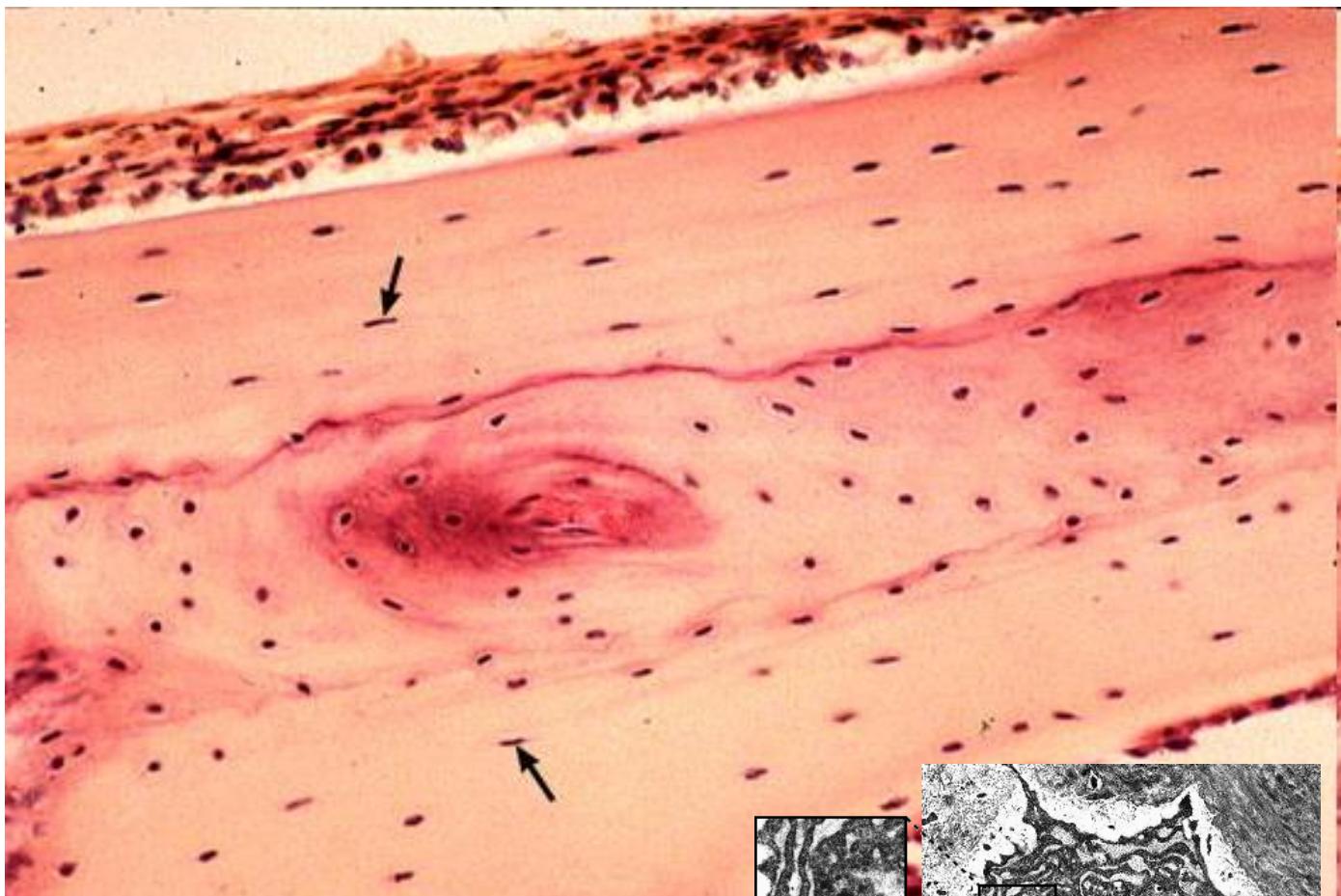
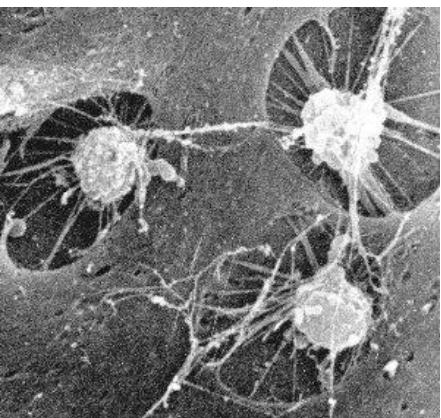


■ Cells of the bone - osteoblasts

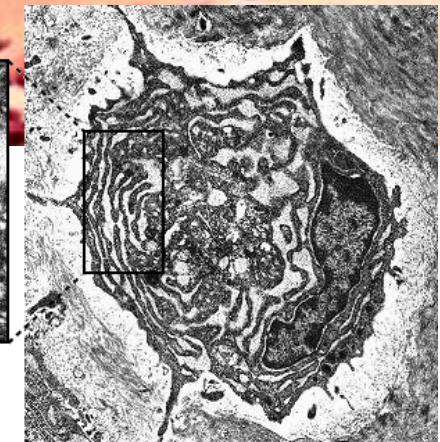
- lining bone surface
 - produce ECM – collagen (I) and noncollagenous proteoglycans, glycoproteins
 - basophilic cytoplasm, rER, well developer Golgi Apparatus
 - euchromatin nucleus
-
- **osteocytes** embedded in matrix
 - *canaliculi ossium*



Cells of the bone - osteocytes

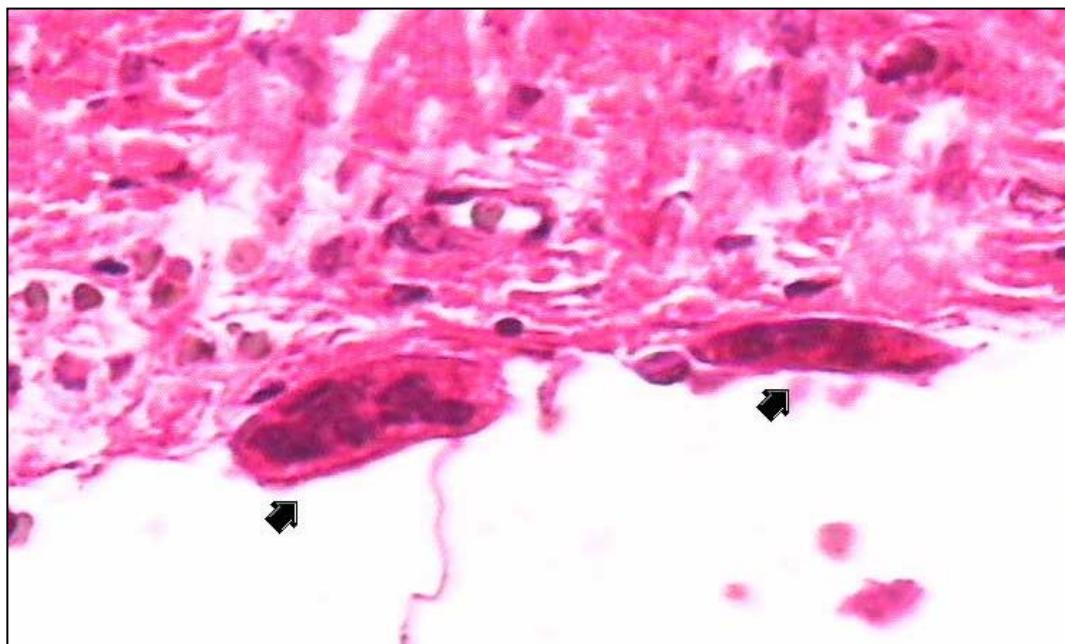
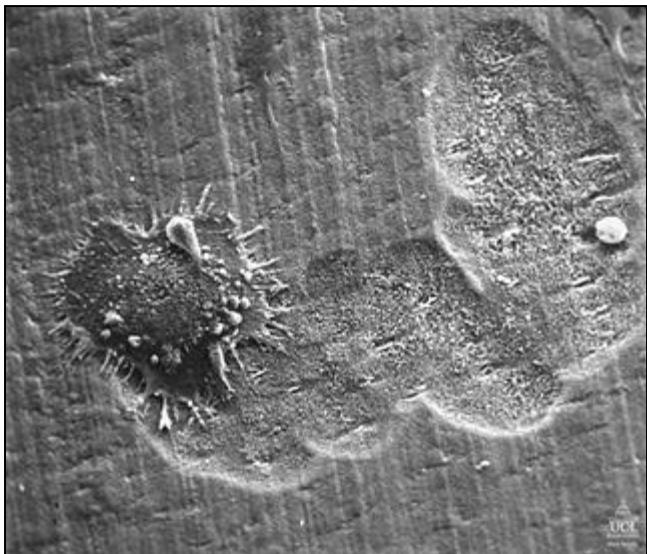


RER
-rough
endoplasmic
reticulum

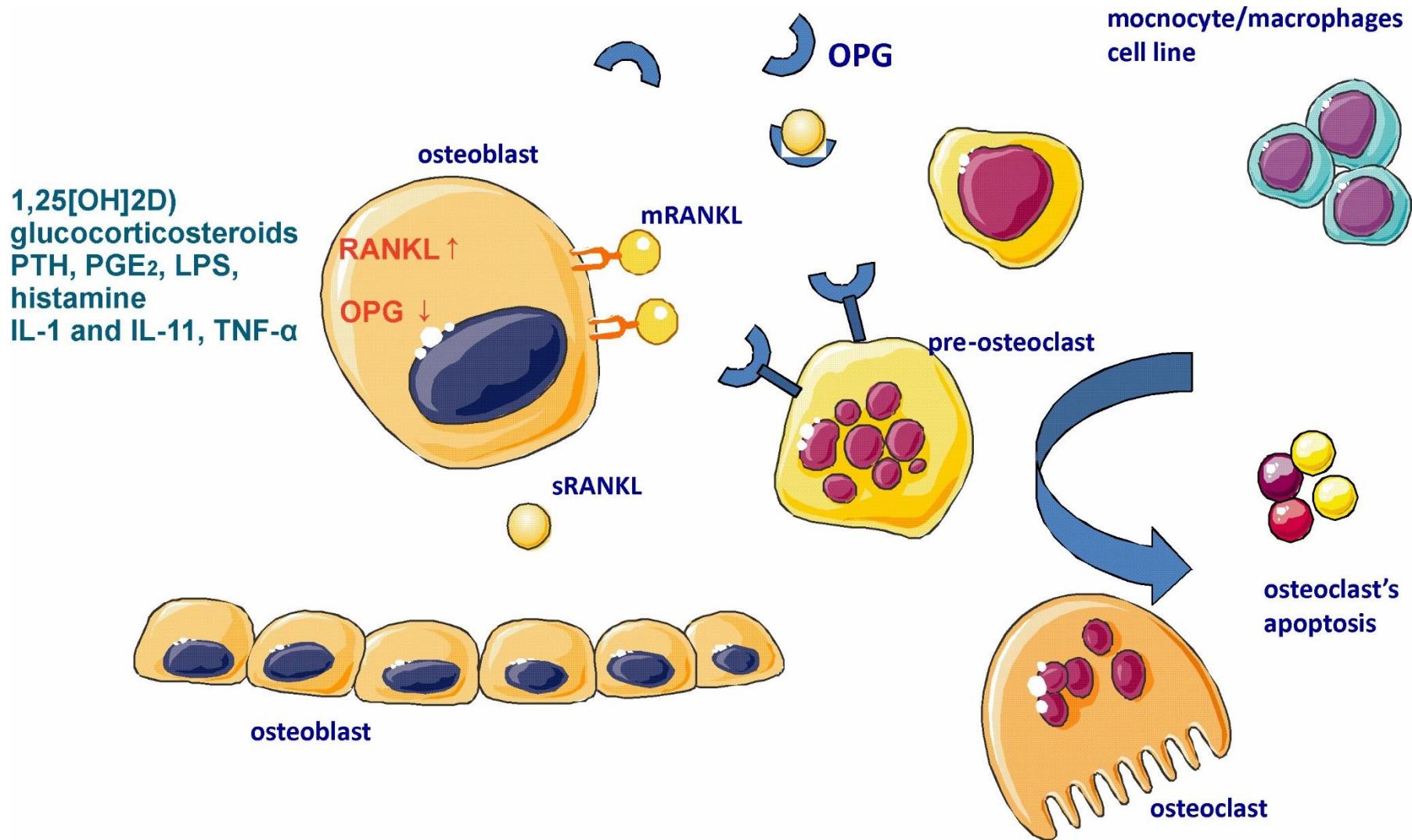


■ Cells of the bone - osteoclasts

- multinuclear, formed by fusion of mononuclear macrophages
- bone matrix resorption

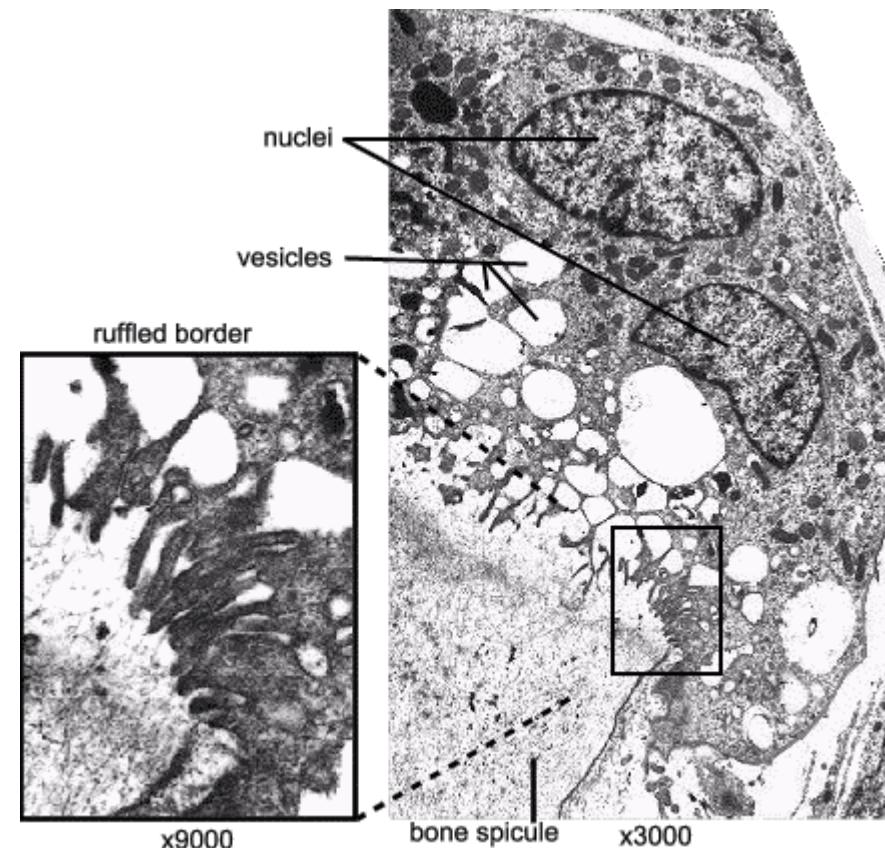
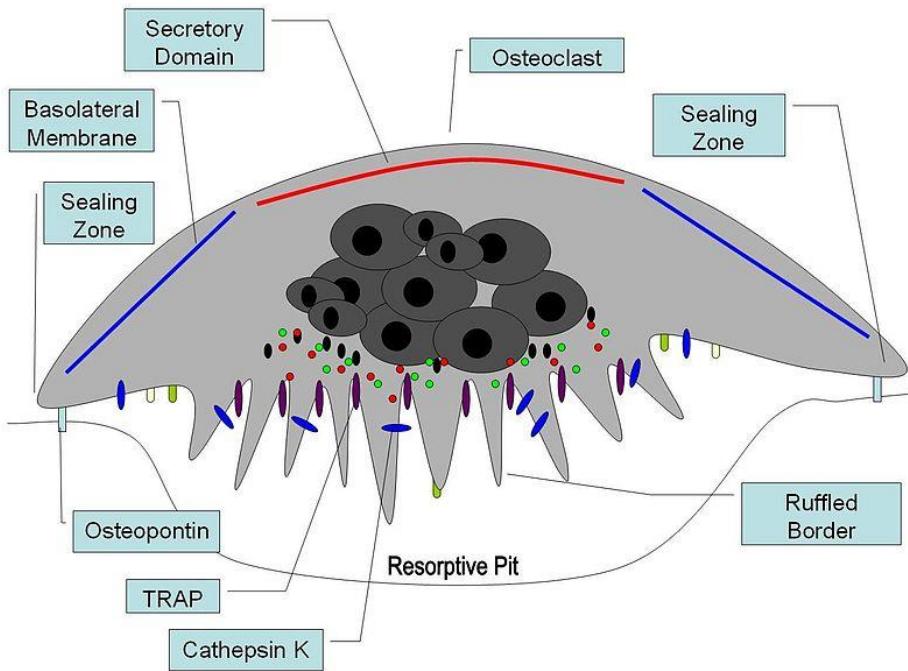


Cells of the bone - osteoclasts



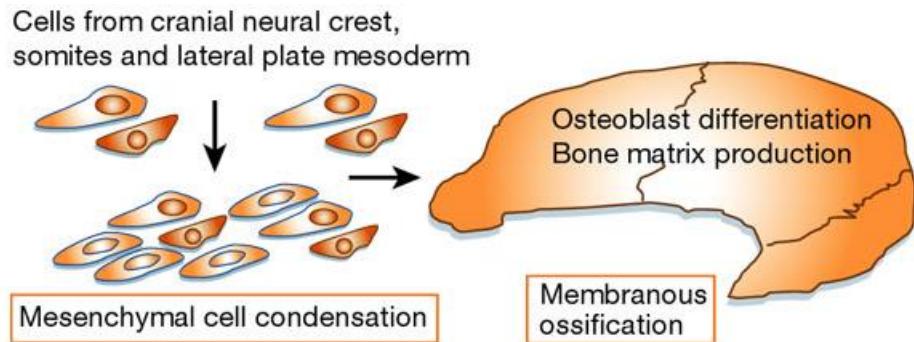
■ Cells of the bone - osteoclasts

- complex architecture
- enzymes degrading organic matrix
- HCl

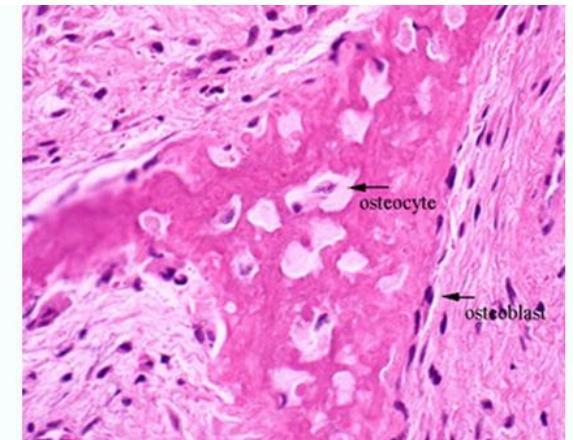
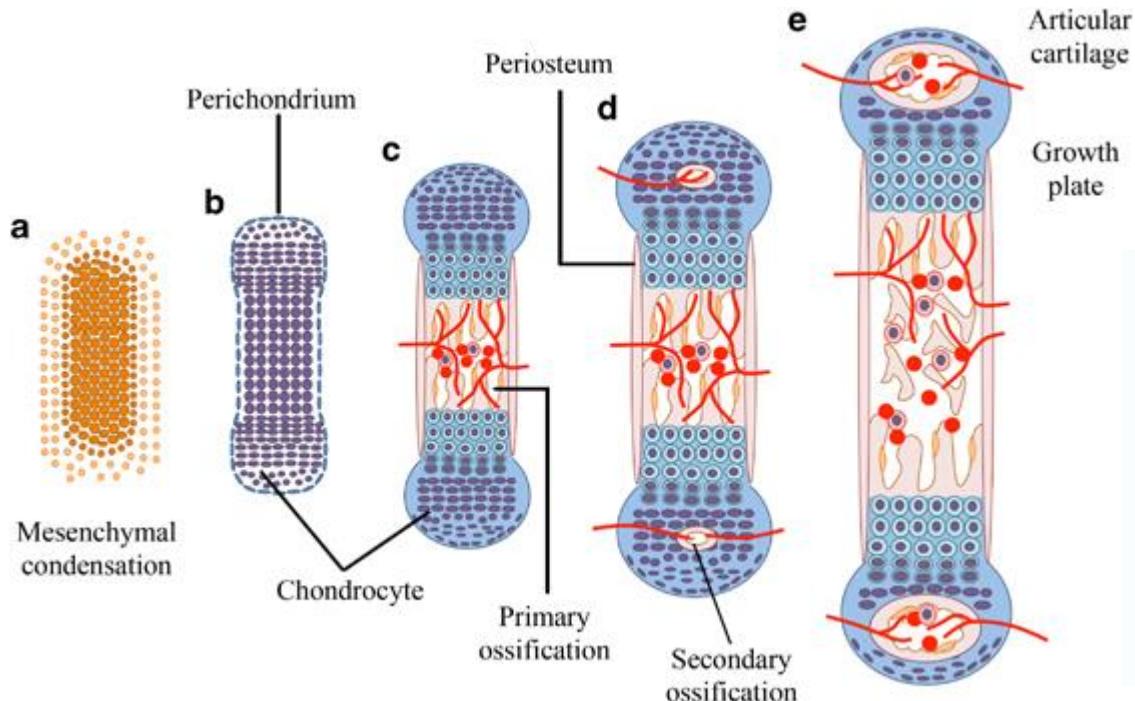


■ Ossification

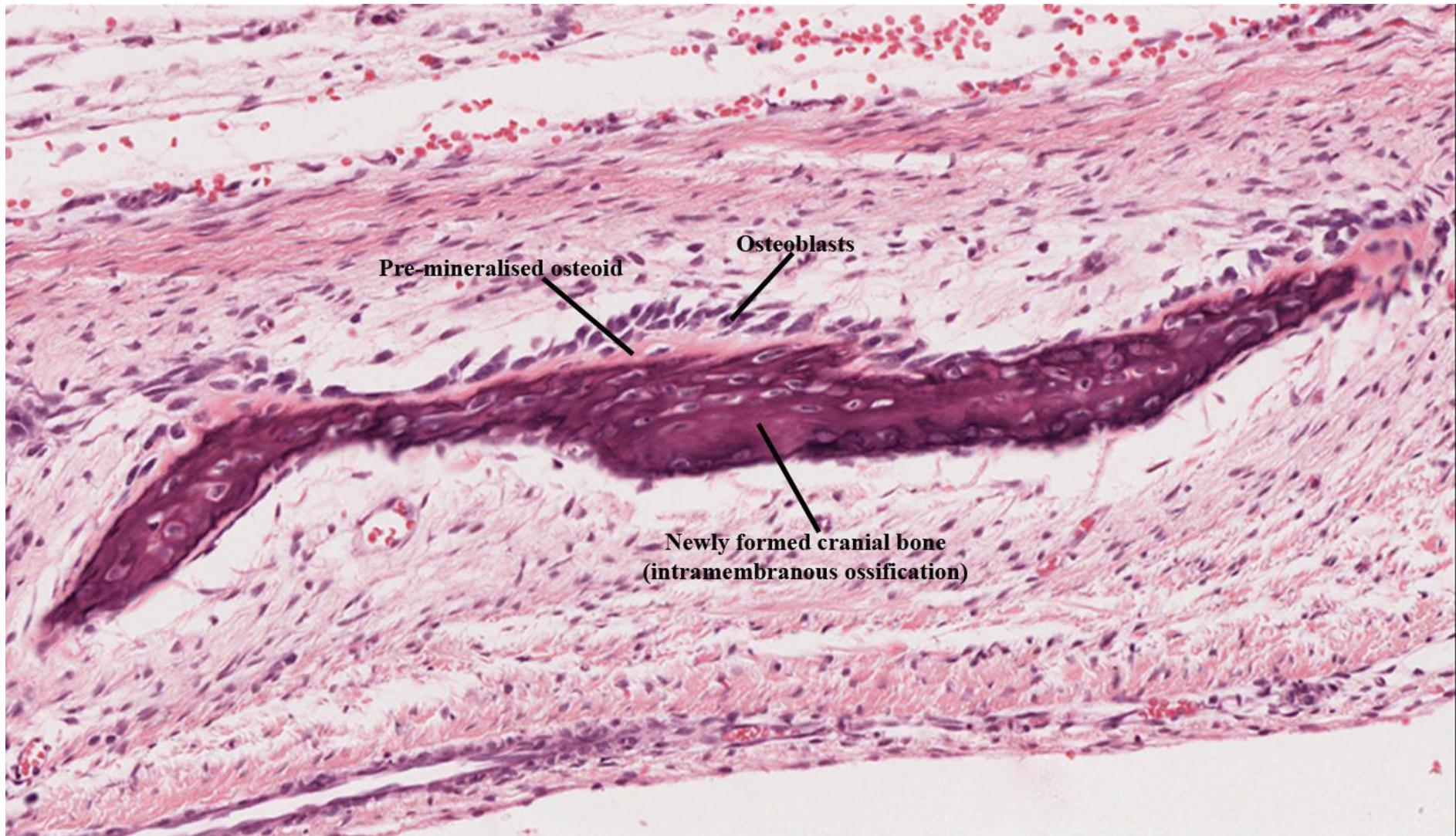
Intramembranous



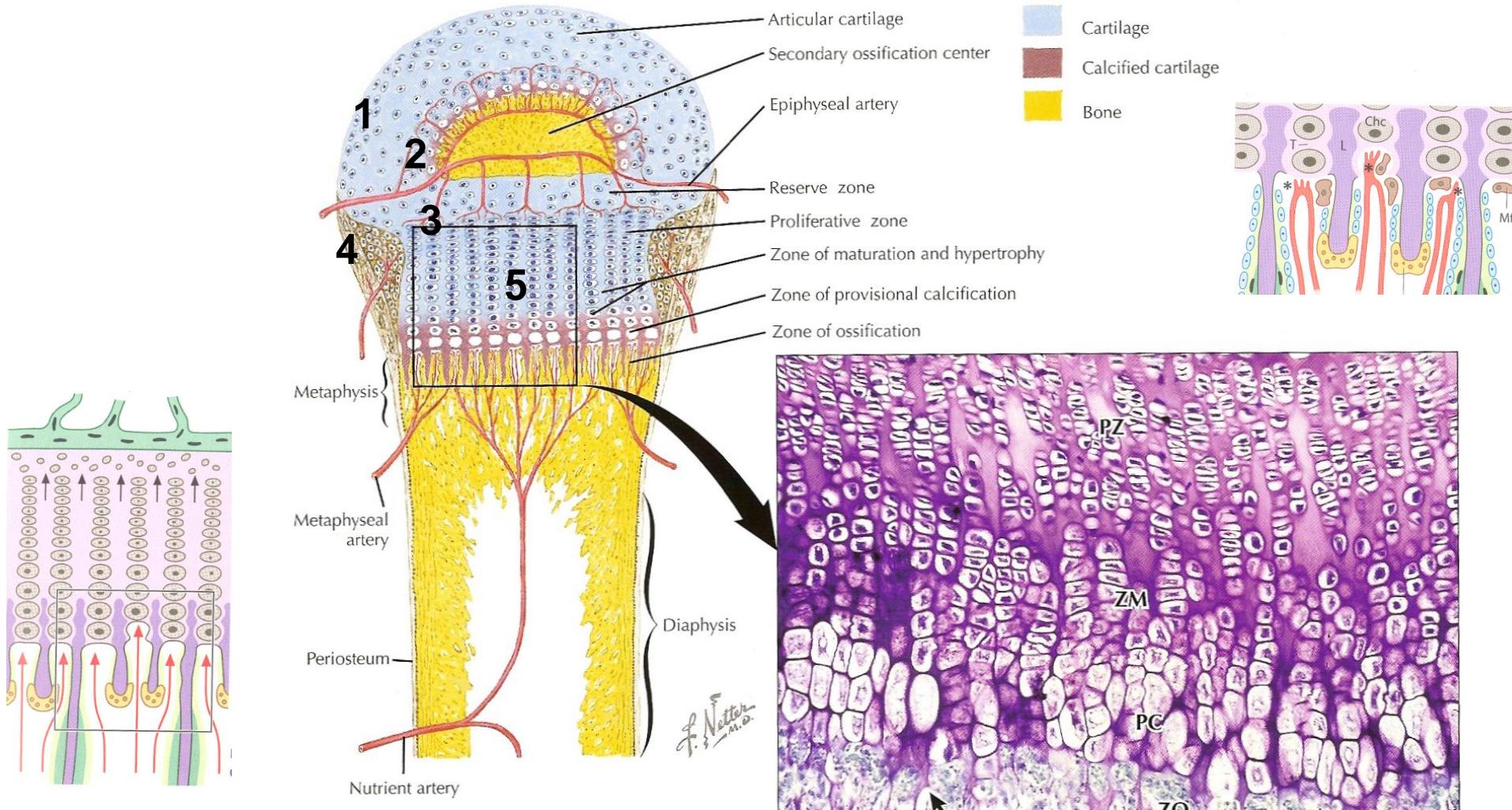
Endochondral



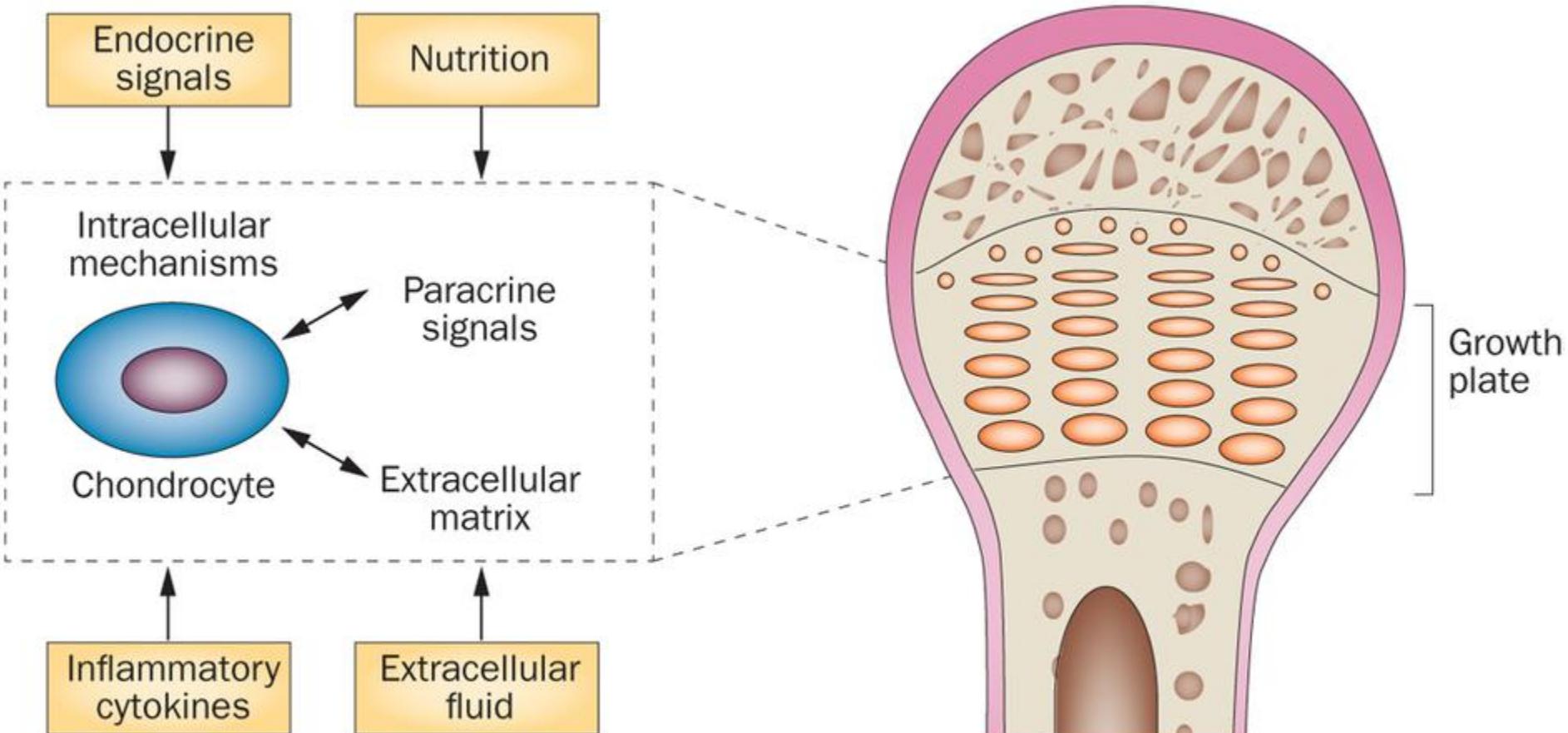
■ Intramembranous ossification



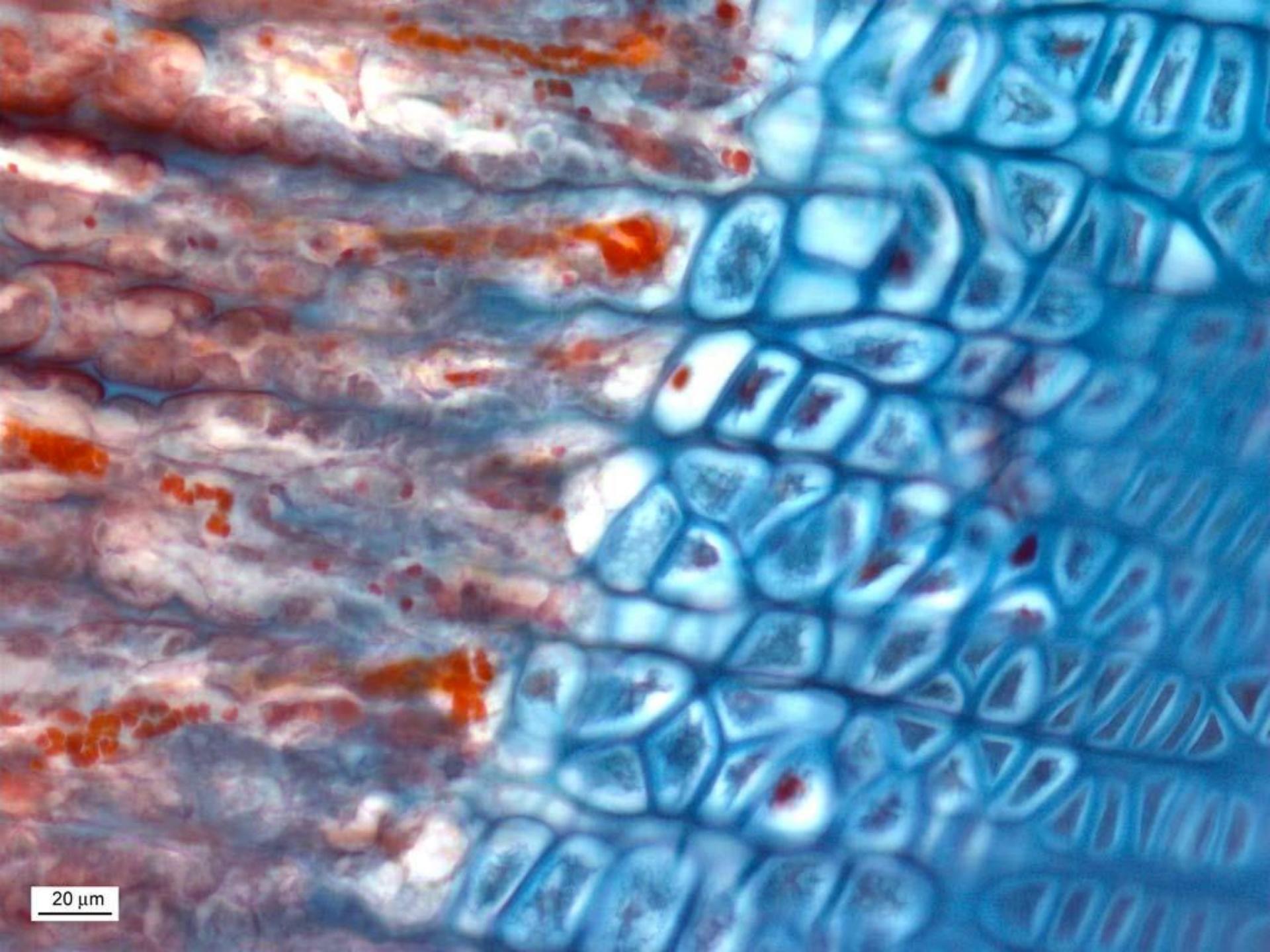
■ Endochondral ossification



■ Endochondral ossification

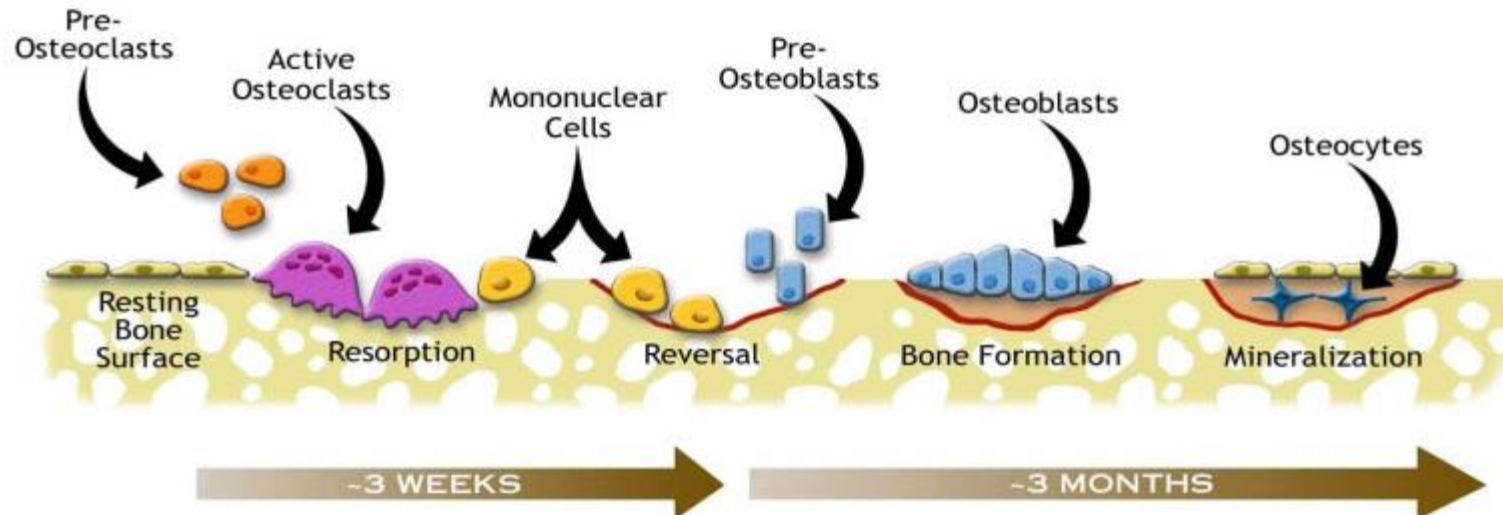




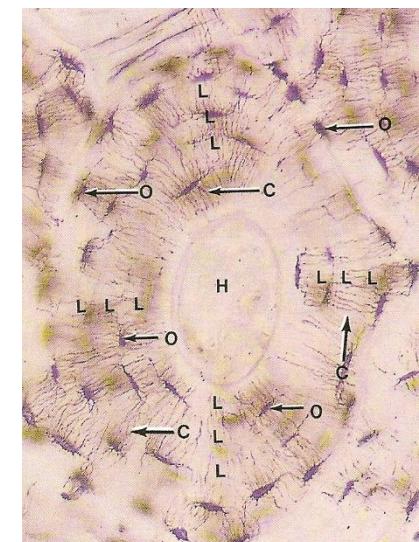


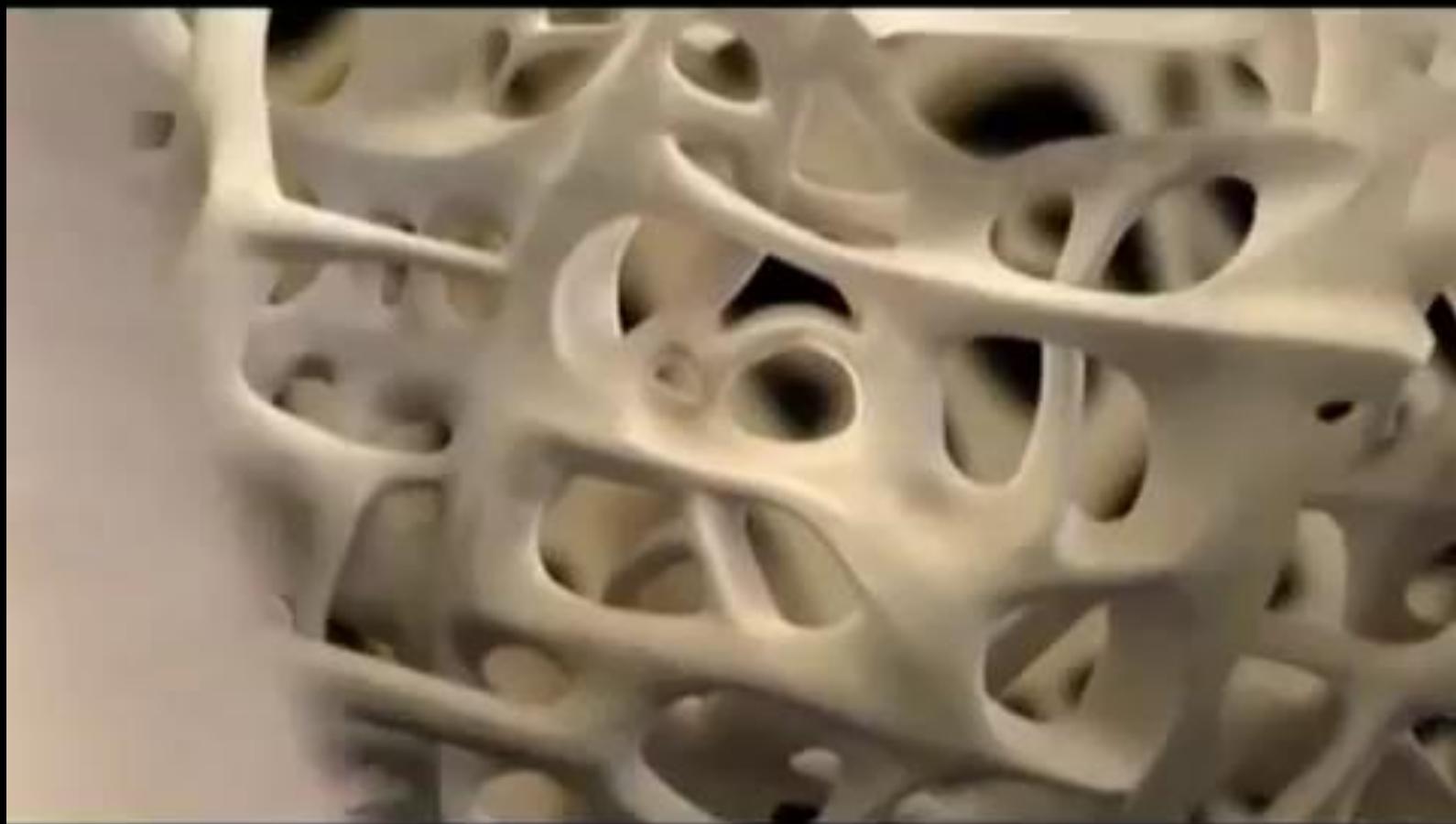
20 μm

■ Bone remodelling

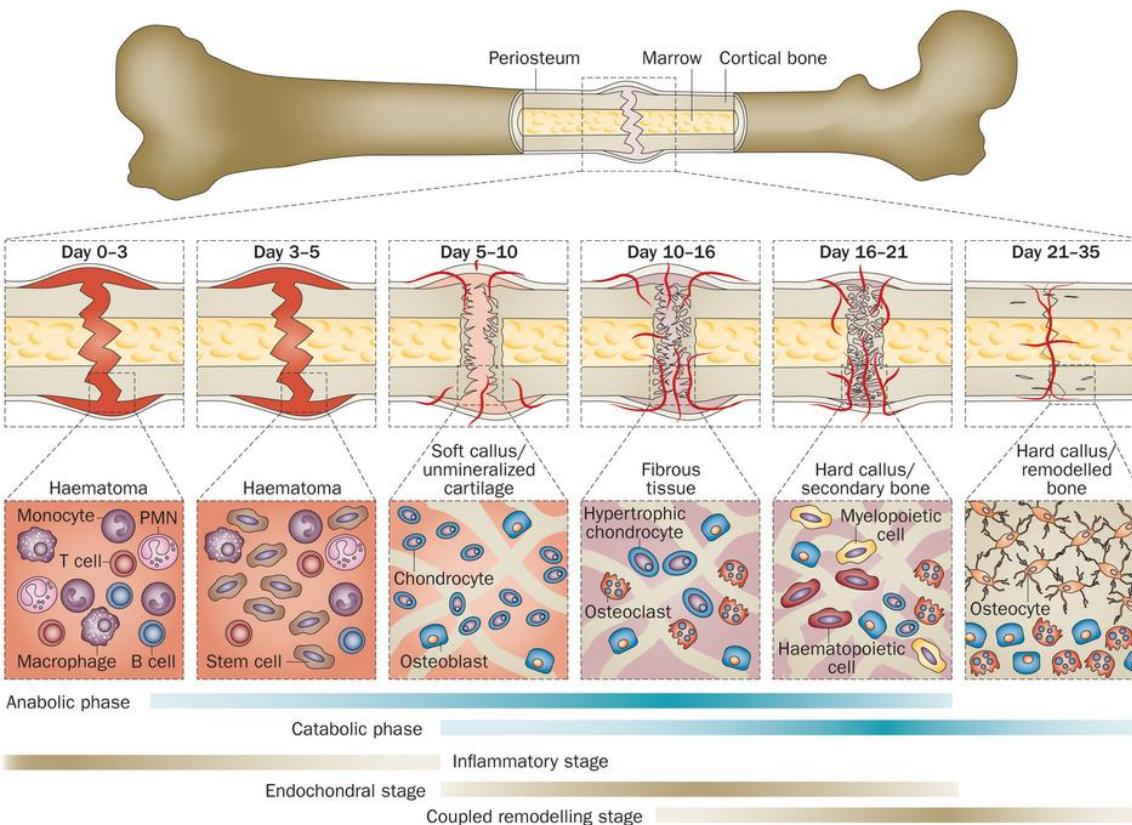


<http://ns.umich.edu/Releases/2005/Feb05/img/bone.jpg>





▪ Clinical correlations - fracture healing



Reactive Phase

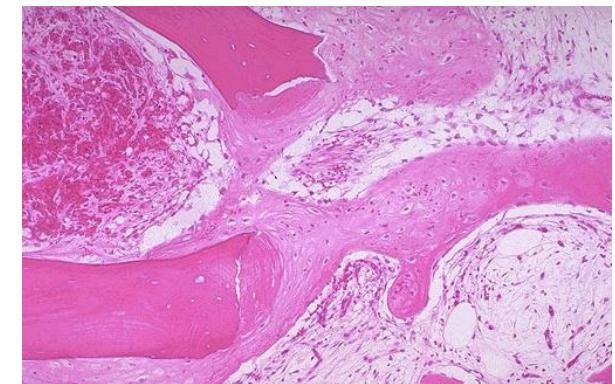
- fracture and inflammatory phase
- granulation tissue formation

Reparative Phase

- cartilage *callus* formation
- lamellar bone deposition

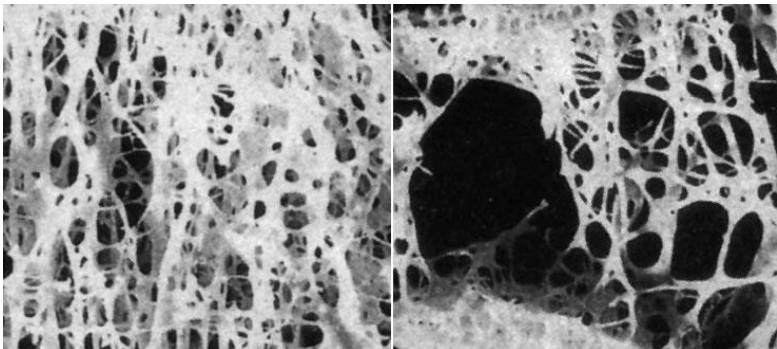
Remodeling Phase

- remodeling to original bone shape



- **Clinical correlations – disbalance in osteosynthesis and osteoresorption**

- **OSTEOPOROSIS**



- **REVMATOID ARTHRITIS**



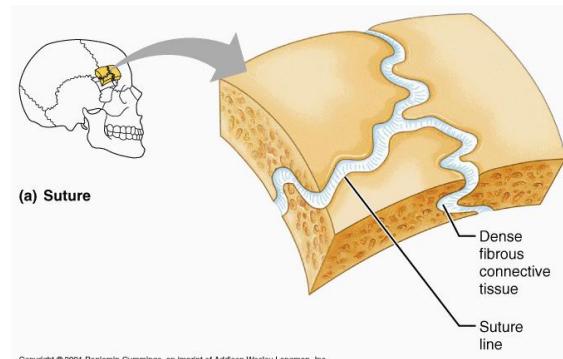
- **OSTEOPETROSIS**



Skeletal joints

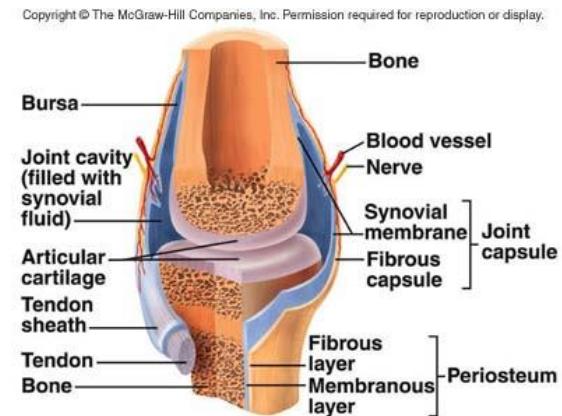
Synarthrosis

- joint by intercalated tissue (cartilage, bone or c.t.)
 - **Synostoses** – joint by bone tissue – os coxae, os sacrum
 - **Synchondrosis** – joint by hyaline cartilage – development of synostosis
 - **Symphysis** – joint by fibrocartilage – os pubis, intervertebral discs
 - **Syndesmosis** – dense collagen regular c.t. – sutures of skull, gomphosis



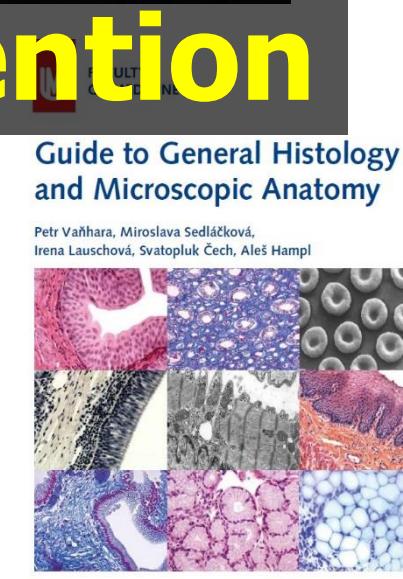
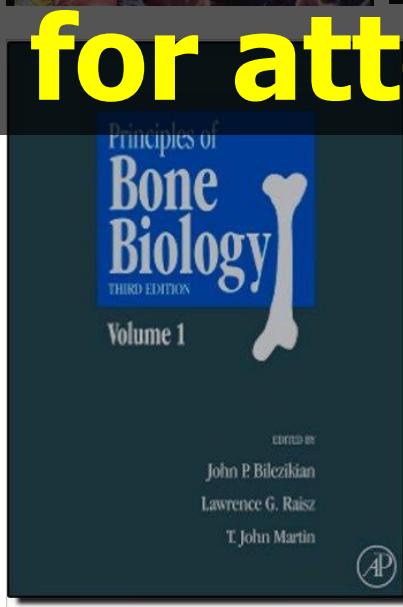
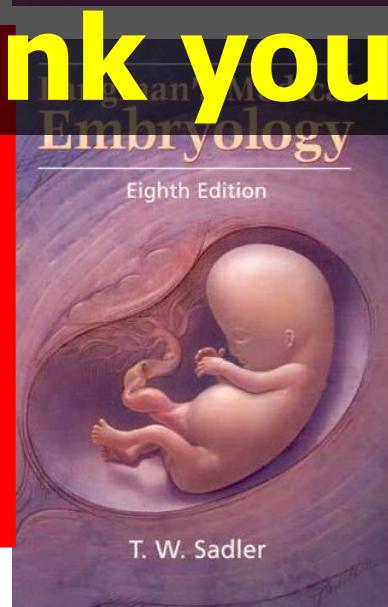
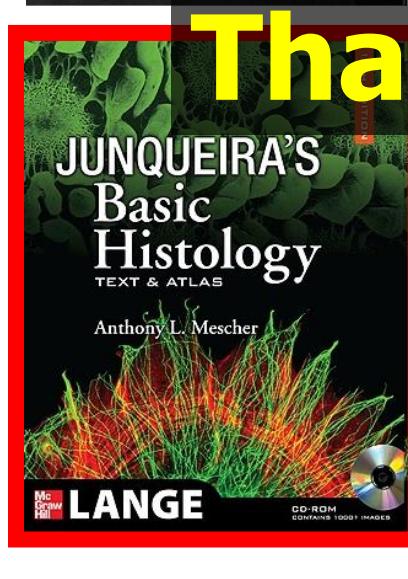
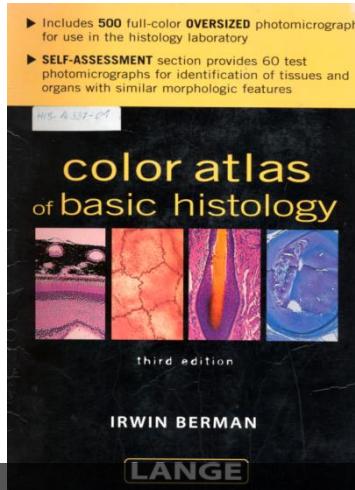
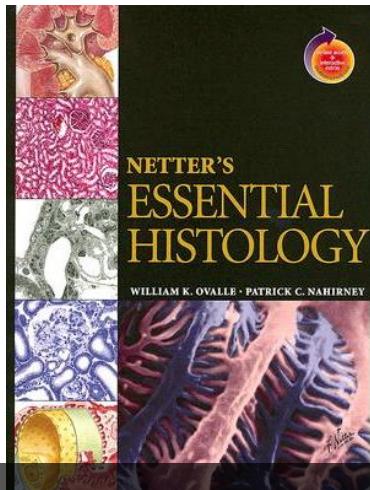
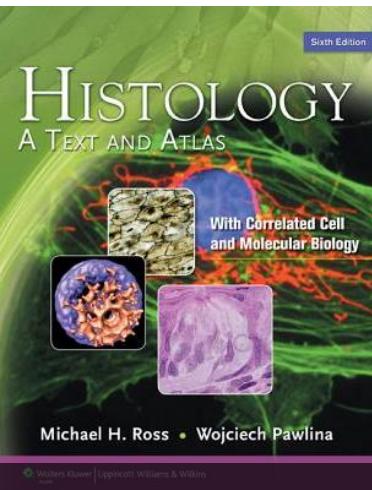
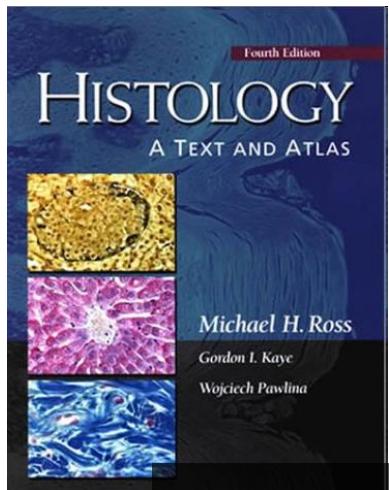
Diarthrosis

- synovial joint
 - hyaline cartilage without perichondrium
 - cartilage calcification in site of attachment to the bone
 - joint capsule
 - *Stratum fibrosum*
 - *Stratum synoviale*



- Meniscus – fibrocartilage, avascular, without innervation
- tendons – dense collagen regular c.t., elastic fibers
- bursae – like joint capsule

Further study



Masaryk University, Brno 2012