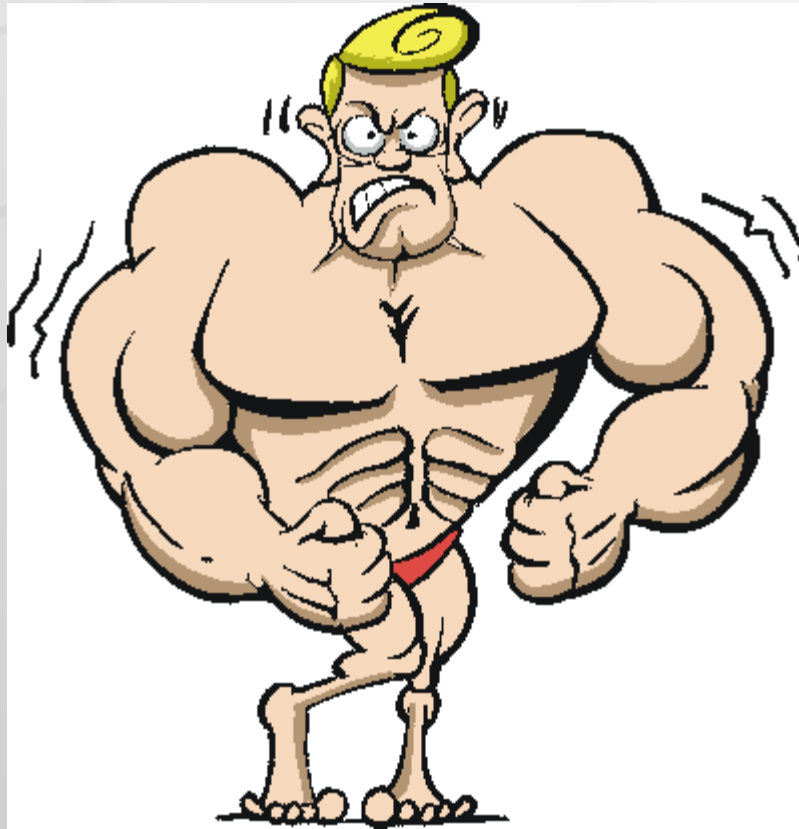




Muscle tissue



- 1) Striated skeletal muscle tissue.
- 2) Striated cardiac muscle tissue.
- 3) Smooth muscle tissue.



General characteristic of muscle tissue

- ✦ Origin: mesoderm and mesenchyme
- ✦ Excitability
- ✦ Contraction + relaxation \Rightarrow cause movement
- ✦ Composition: **muscle cells** + **connective tissue** (+blood vessels + nerves)
- ✦ contractile proteins in sarcoplasm – **actin** and **myosin**
- ✦ Long axis of cells is usually oriented parallelly with direction of contraction

Nomenclature

mys/myos (muscle)

myocyte (muscle cell)

sarx/sarcós (meat):

cell membrane = **sarcolemma**

cytoplasm = **sarcoplasm**

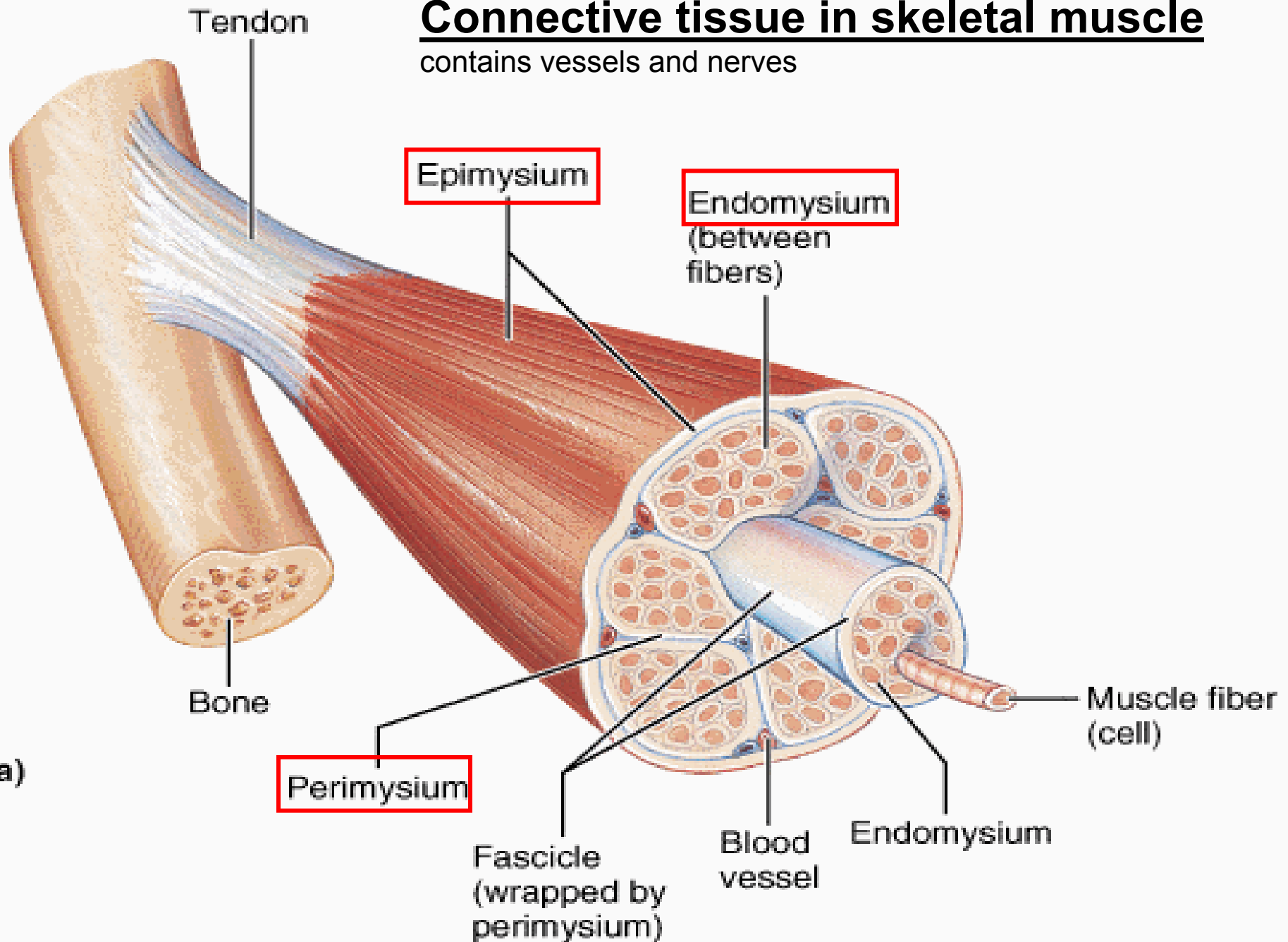
smooth ER = **sarcoplasmic reticulum**

Connective tissue of muscle

- ✦ **Endomysium** – around each muscle cell (fiber)
- ✦ **Perimysium** – around and among the primary bundles of muscle cells
- ✦ **Epimysium** – connective tissue „capsule“ covering the surface of muscle

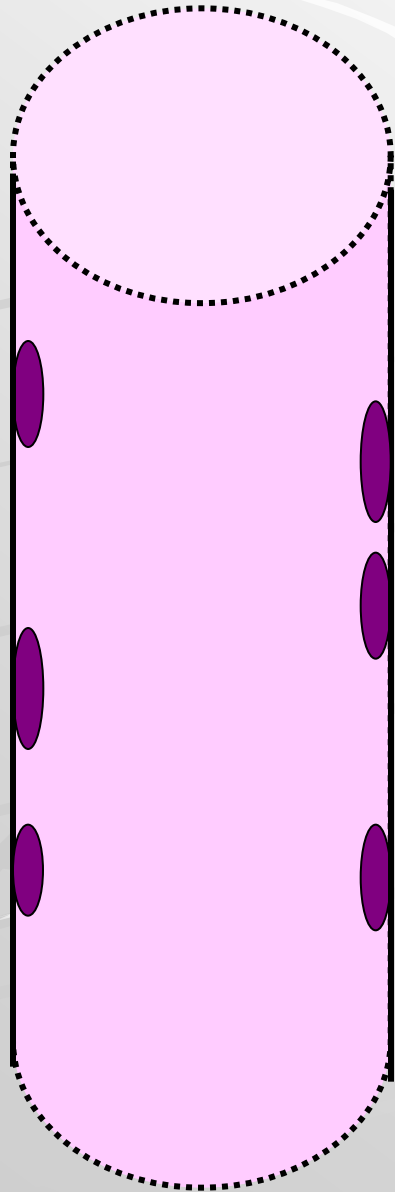
Connective tissue in skeletal muscle

contains vessels and nerves



(a)

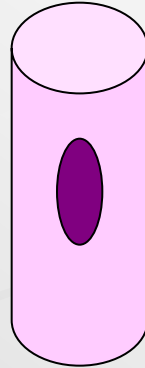
rhabdomyocyte



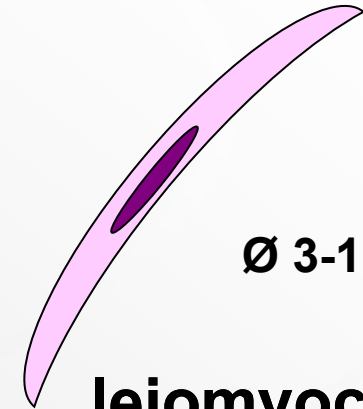
$\text{Ø } 25\text{-}100 \mu\text{m}$

MUSCLE CELLS

cardiomyocyte



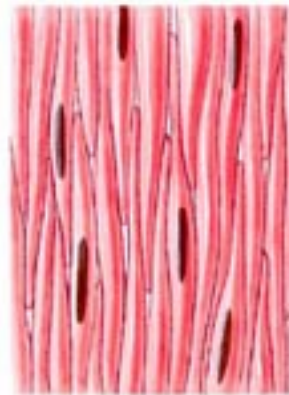
$\text{Ø } 15 \mu\text{m}$



$\text{Ø } 3\text{-}10 \mu\text{m}$

leiomyocyte

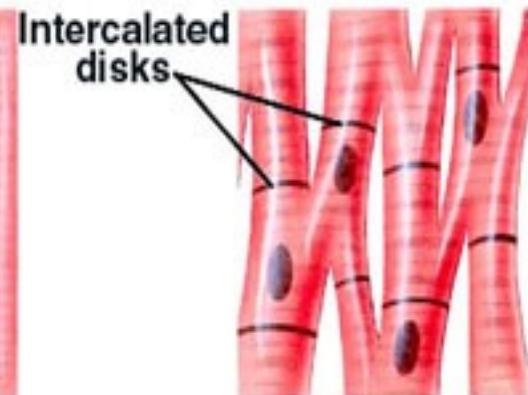
Types of Muscle



Smooth muscle



Skeletal muscle



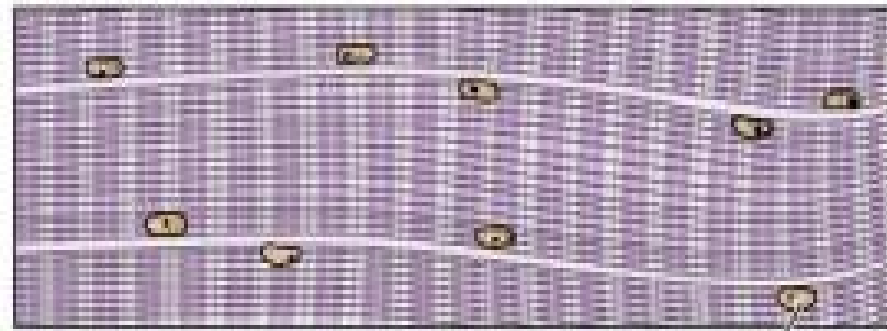
Cardiac muscle

Muscle types

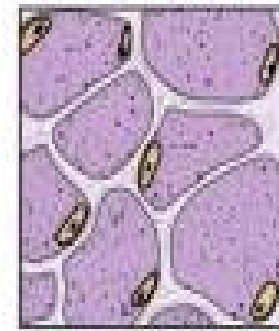
Activity

Occurrence:

Skeletal muscle



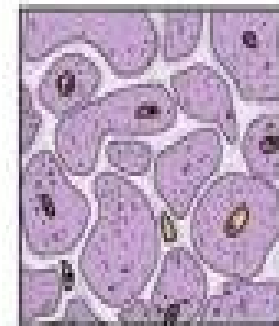
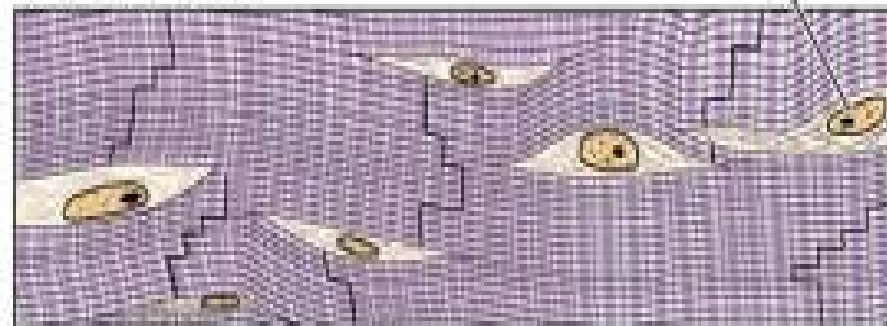
Cross sections



Strong, quick discontinuous voluntary contraction

Skeletal muscles

Cardiac muscle

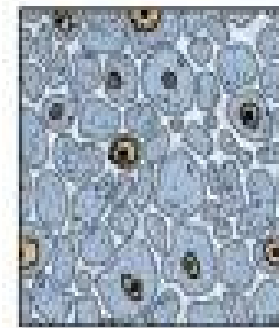
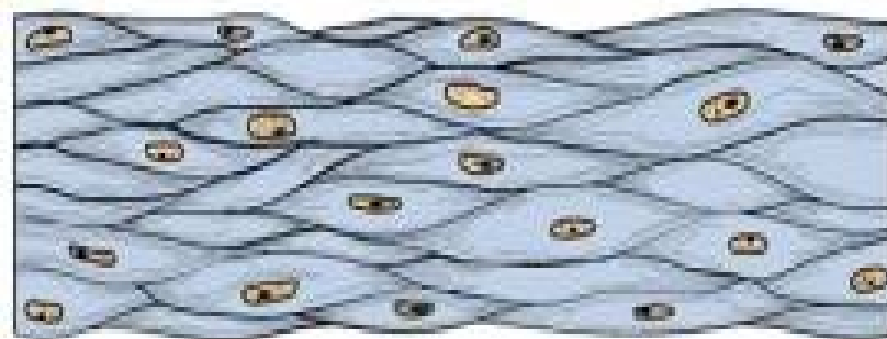


Strong, quick continuous involuntary contraction

Heart

Smooth muscle

Intercalated disks

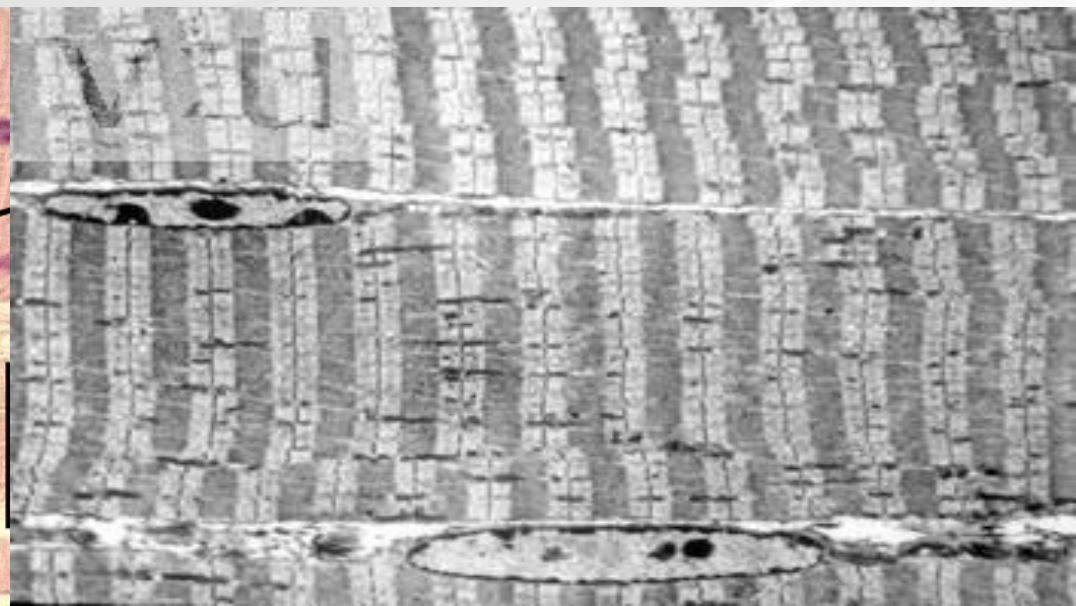
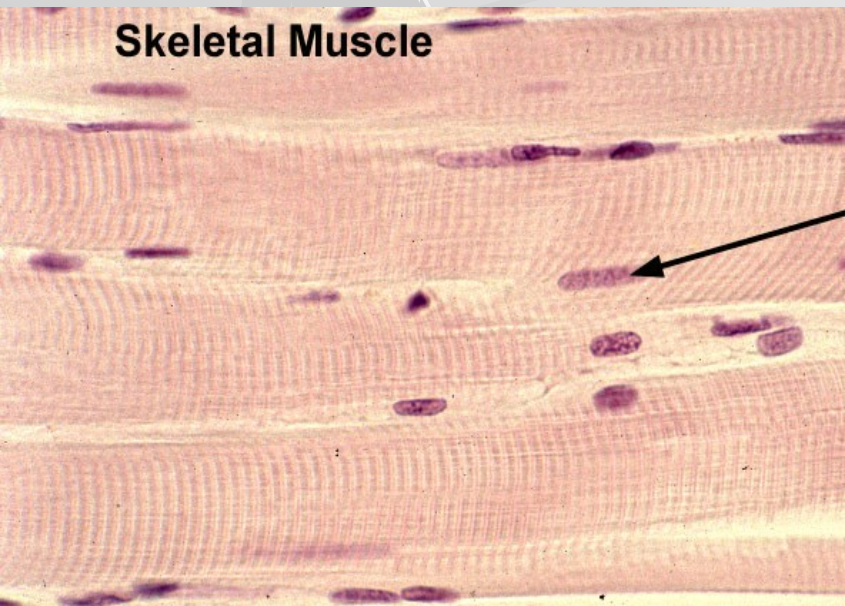


Weak, slow involuntary contraction

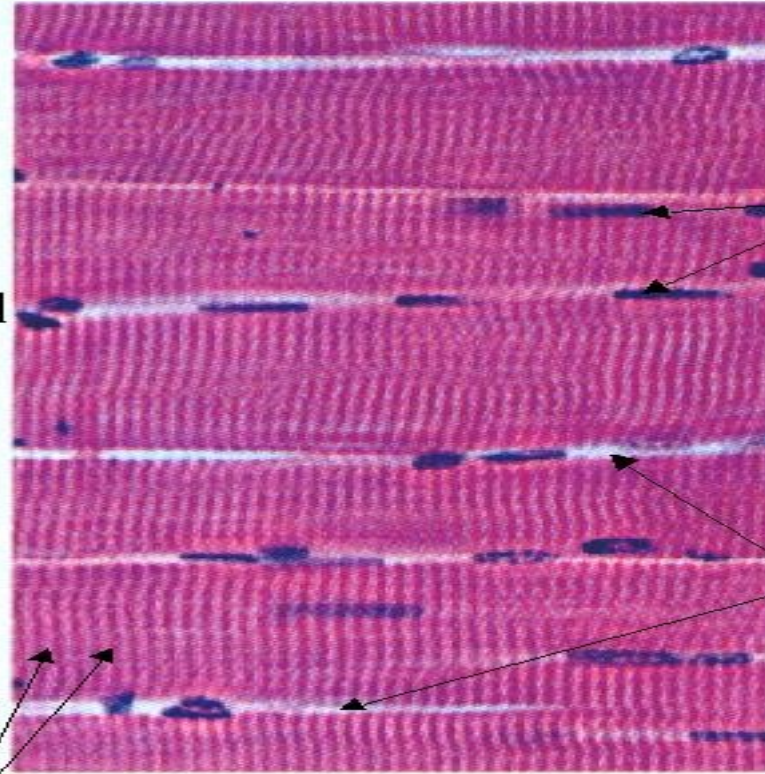
Inner organs (their wall)

Cross-striated skeletal muscle tissue

- ✦ **morphological and functional unit: muscle fiber (rhabdomyocyte)** – elongated, cylindrical shape, multinucleated cell (=syncytium) – nuclei are located at the periphery (beneath sarcolemma), myofibrils show cross striation
- ✦ diameter of muscle fiber: 25-100 μm
- ✦ length: millimeters - centimeters (up 15)



Skeletal Muscle Characteristics



nuclei

Connective endomysium separates cells.

Myofibrils fill cell interior



Skeletal muscle cells are long multinucleated cylinders, separated by connective tissue.

Striations are the dark bands perpendicular to cell length

Structure of rhabdomyocyte

■ **sarcolemma + T-tubules**

■ **nuclei** (25-40 per 1mm of the length)

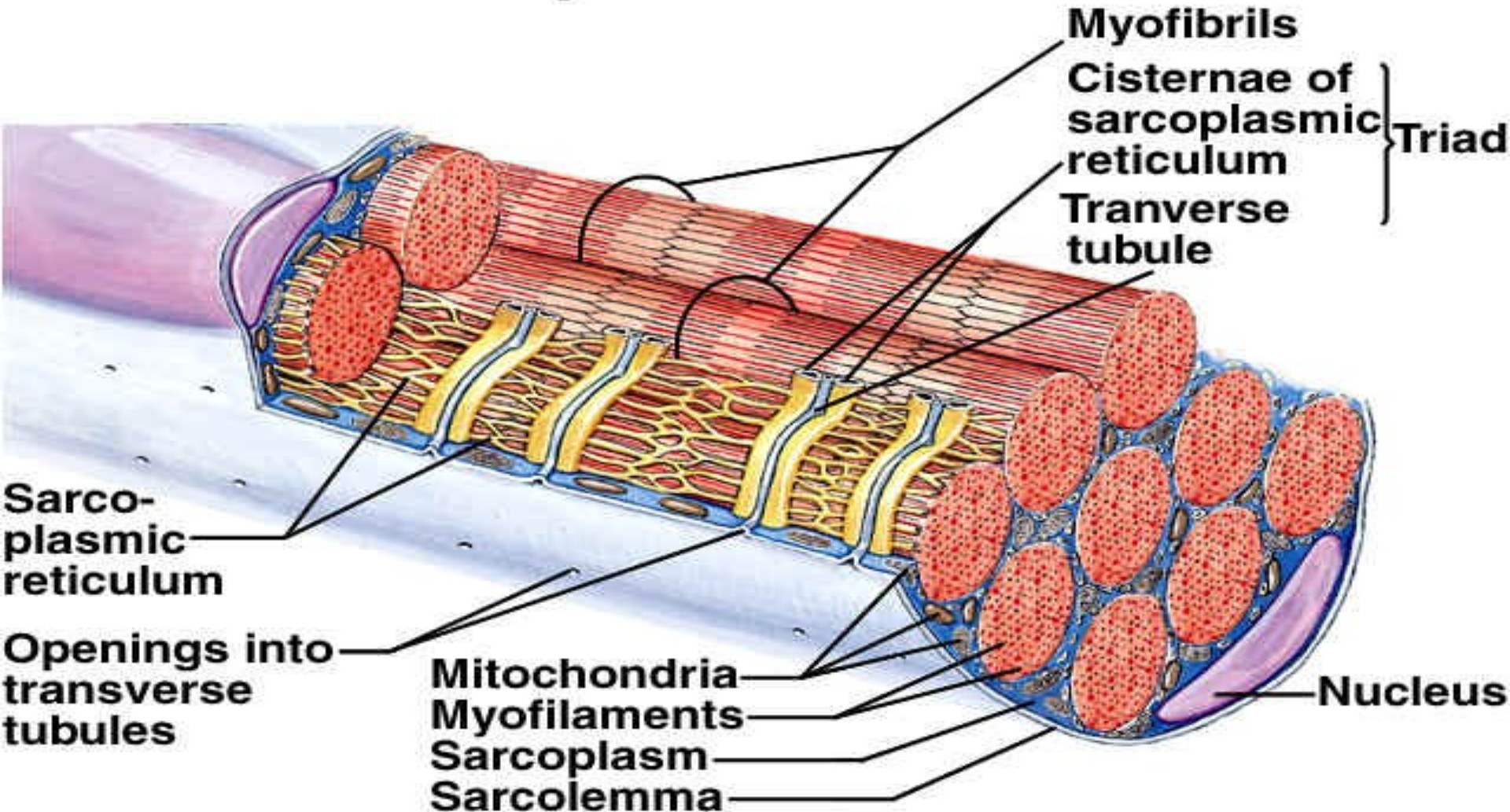
■ **sarcoplasm:**

- **myoglobin** (protein with heme ([iron](#)-containing [porphyrin](#)) prosthetic group, and is the primary [oxygen](#)-carrying [pigment](#))
- **myofibrils** (cross-striated 1–2 μm thick fibrils)
- **organelles: mitochondria, Golgi apparatus, sarcoplasmic reticulum** (Ca^{2+} depot , sarcotubules, terminal cisternae)
- **inclusion (glycogen)**

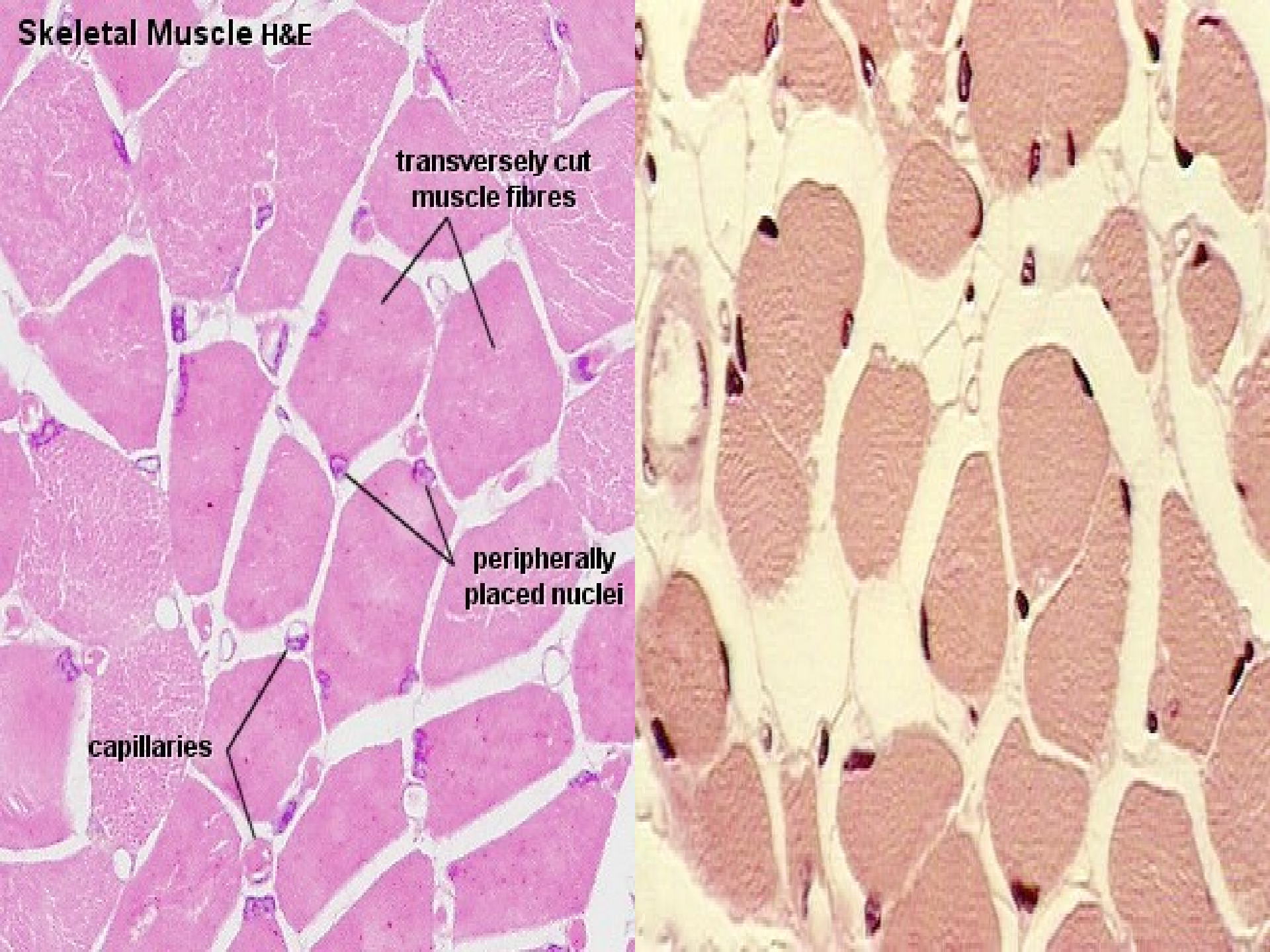
Rhabdomyocyte

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Sarcoplasm Contents



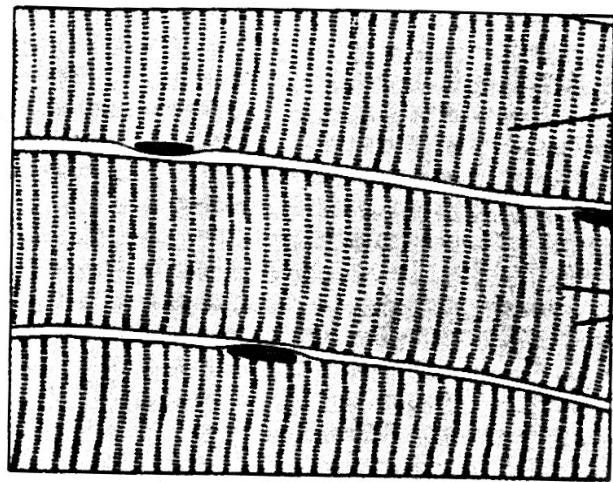
Skeletal Muscle H&E



**transversely cut
muscle fibres**

**peripherally
placed nuclei**

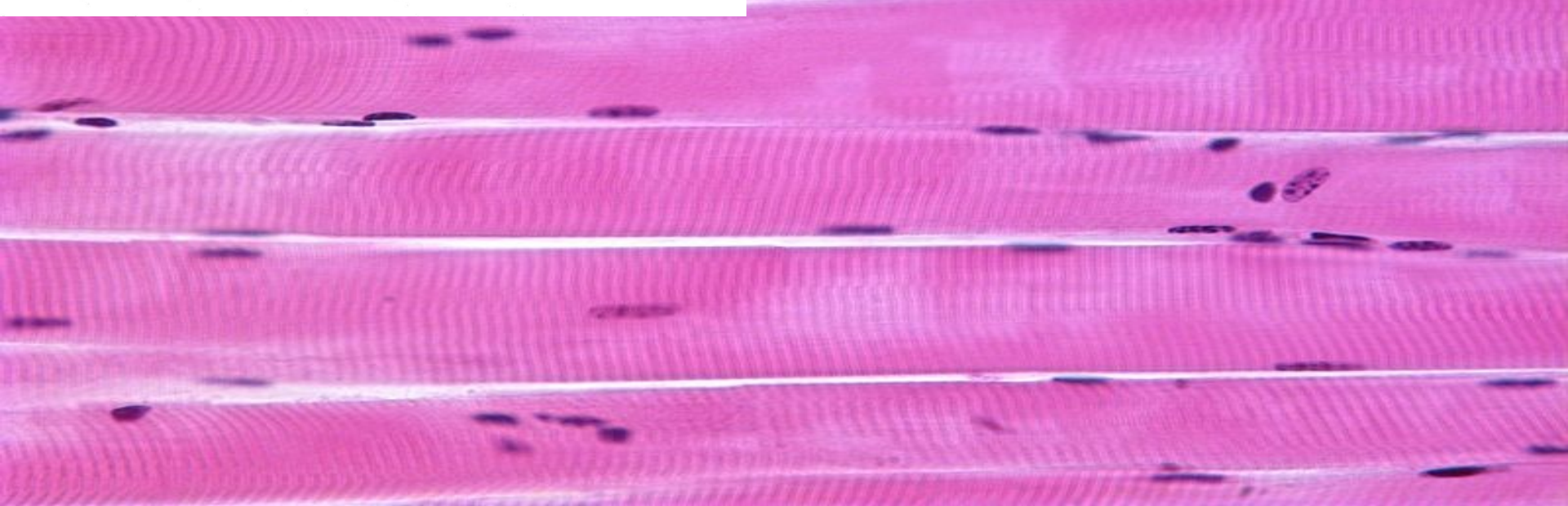
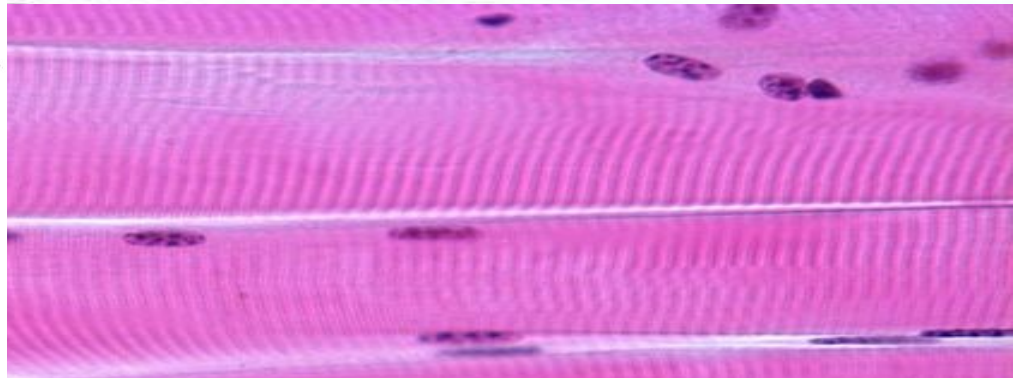
capillaries



Muscle cell

Nucleus

Striations



Skeletal muscle cell (fiber)

< rhabdomyocyte >

Used terms:

Muscle fiber = myofiber = syncitium = rhabdomyocyte

Muscle fiber – morphologic and functional unit of skeletal muscle [Ø 25 – 100 µ]

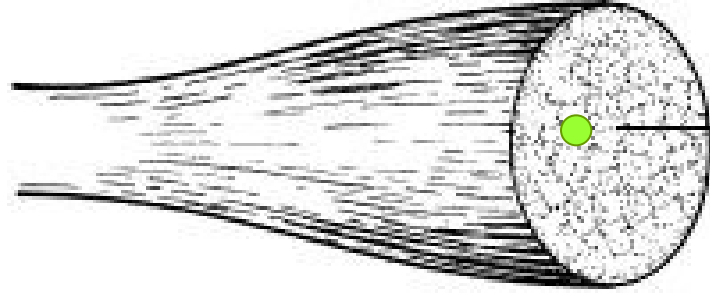
Myofibrils – compartment of fiber sarcoplasm [Ø 0.5 – 1.5 µ]

Myofilaments – actin and myosin, are organized into sarcomeres (several in the length of myofibril) [Ø 8 and 15 nm]

Sarcomere – the smallest contractile unit [2.5 µm in length]

(a)

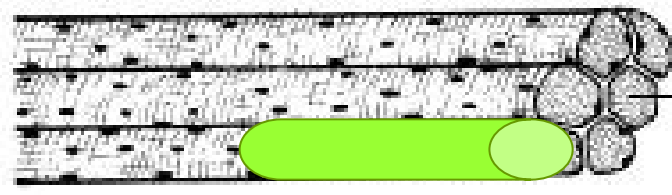
Muscle



Bundles of muscle fibers

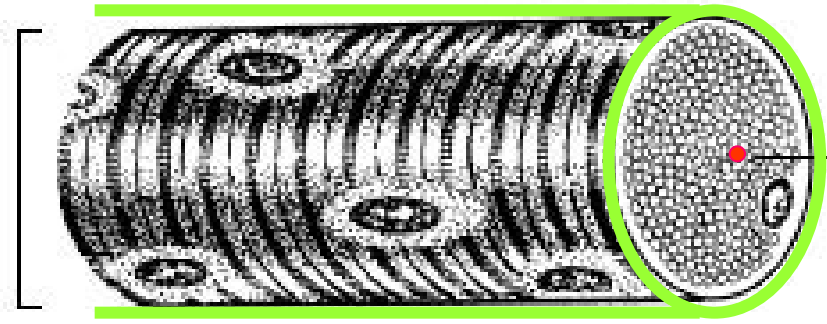
(b)

Bundle of fibres



(c)

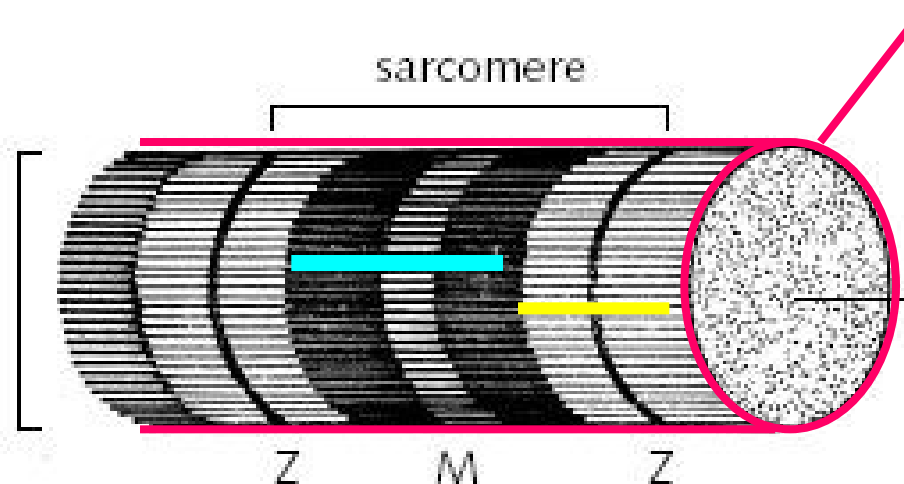
Fibre
50 μm



Muscle fiber – morphologic and functional unit of skeletal muscle
[Ø 25 – 100 μ]

(d)

Myofibril
1 μm



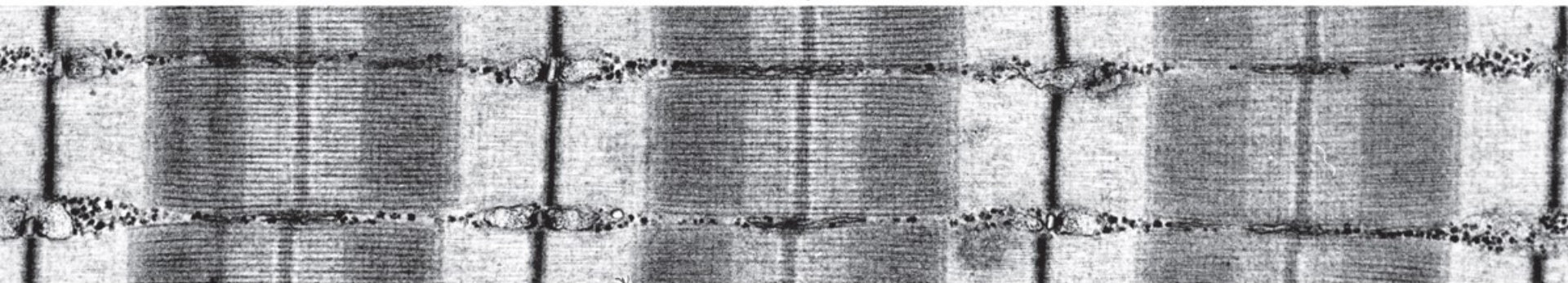
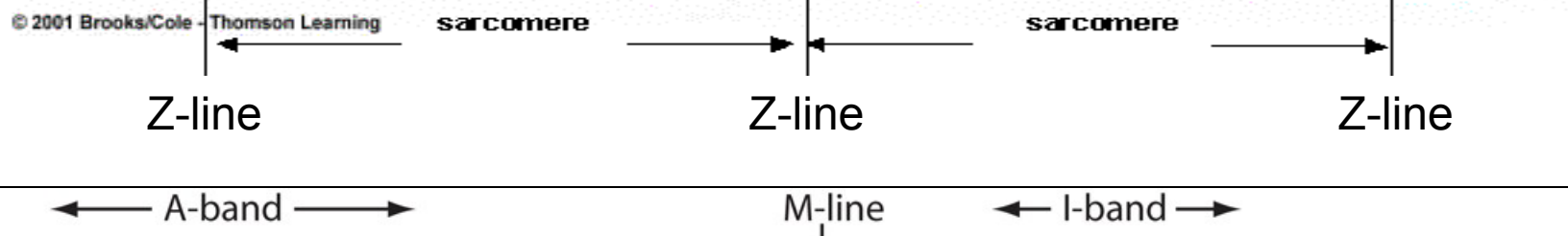
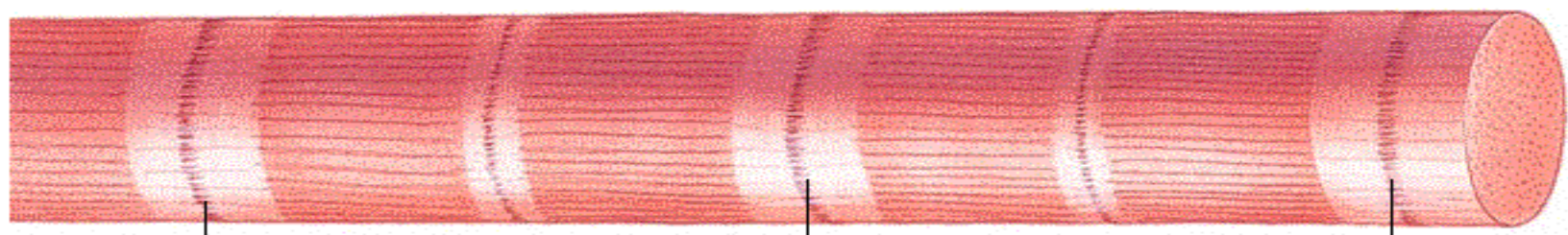
Myofibrils – compartment of fiber sarcoplasm
[Ø 0.5 – 1.5 μ]

Myofilaments – **actin + myosin** are organized into sarcomeres (several in the length of myofibril)
[Ø 8 and 15 nm]

Myofibril [\varnothing 0.5 – 1.5 μ]: elongated structure in sarcoplasm, oriented parallel to the length of cell, composed of 2 types of myofilaments – **actin and myosin** – arranged into:

Sarcomere: the smallest contractile unit between Z-lines on myofibril

(organization of myofilaments causes cross striation of myofibrils)



Z-disc ← Sarcomere → Z-disc

Myofibril structure

isotropic parts (**I-bands**) – light, Z-line (telophragma)

anisotropic parts (**A-bands**) – dark, M-line (mesophragma), H-band

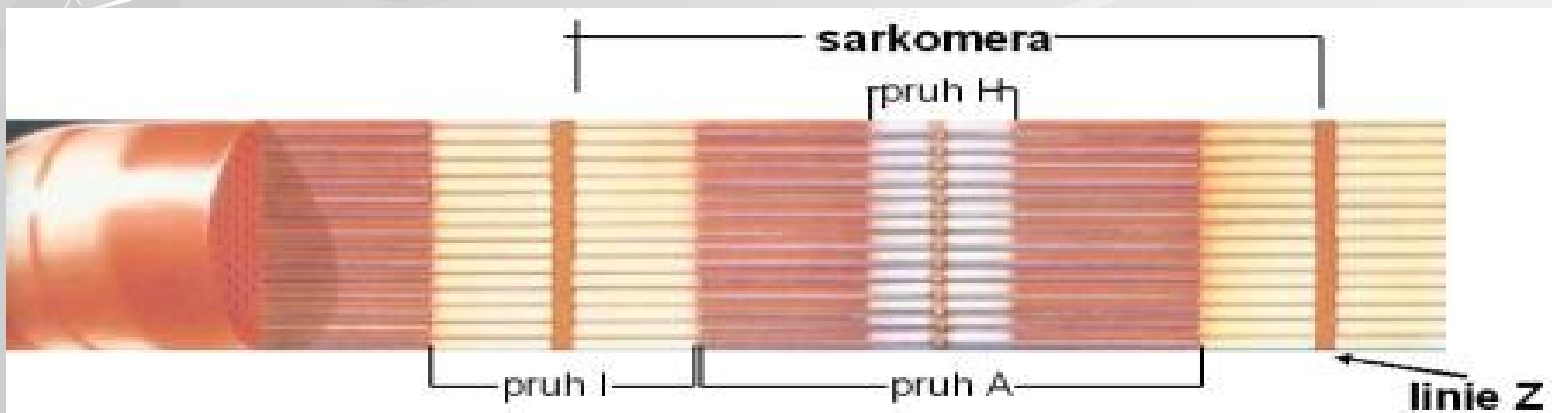
Myofilaments

Thick: only in A-band, **myosin**, 15 nm thick, 1.6–1.8 μm long

Thin: in I-band and at periphery of A-band), **F-actin** and **regulatory proteins** (troponin, tropomyosin), 6 nm thick, 1.5 μm long

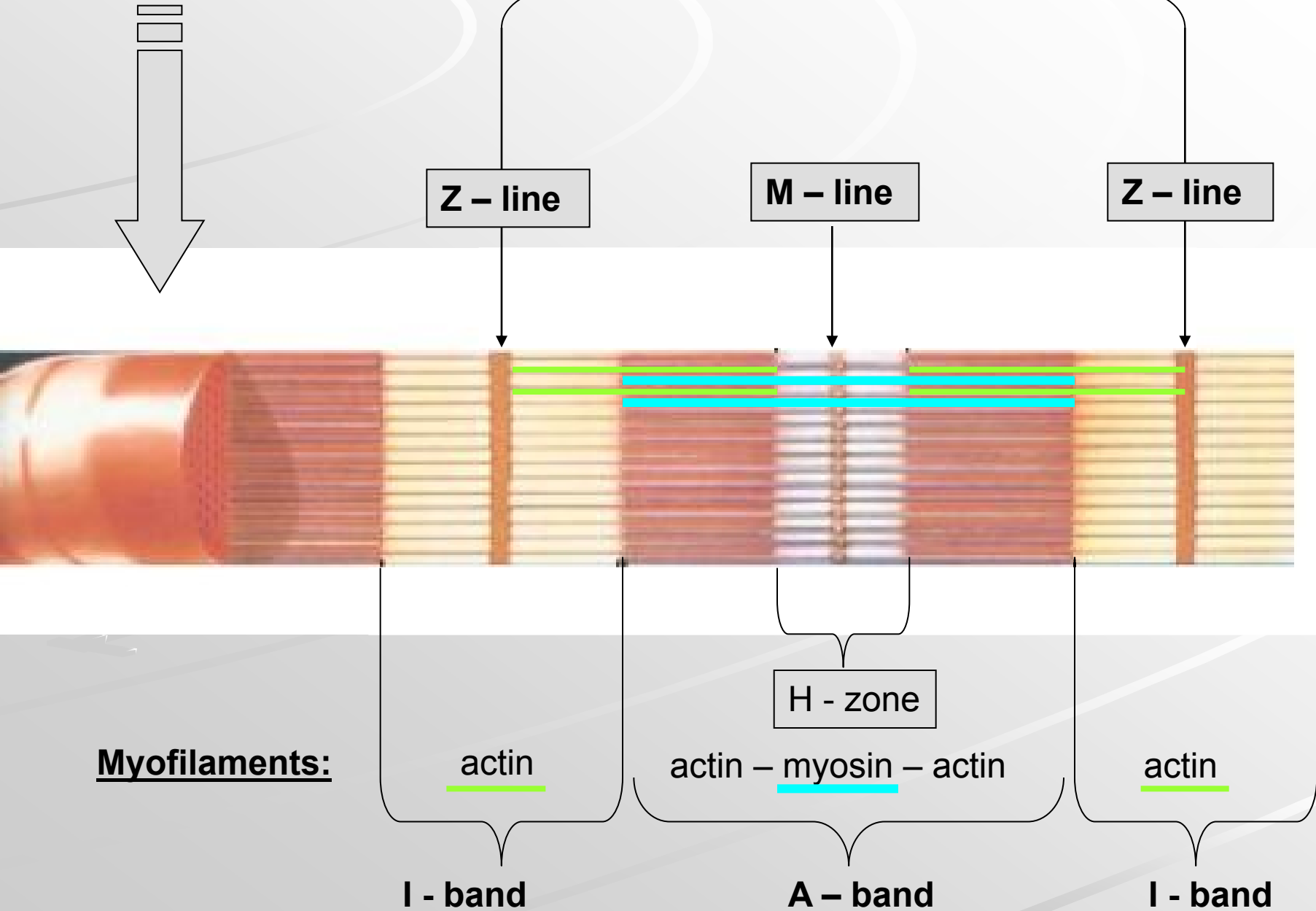
Principle of myofibril shortening during muscle contraction: sliding mechanism mechanismu

the length of myofilaments does not change during contraction



MYOFIBRIL

SARCOMERE



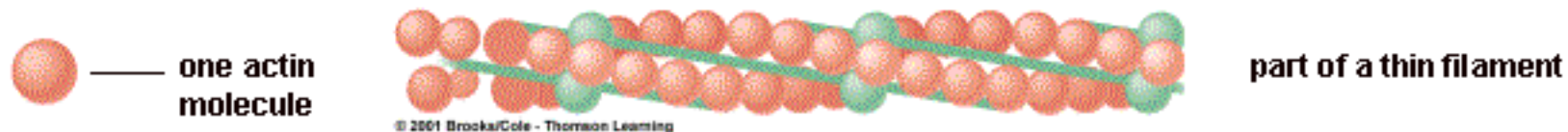
Muscle Microfilaments

Thin filaments

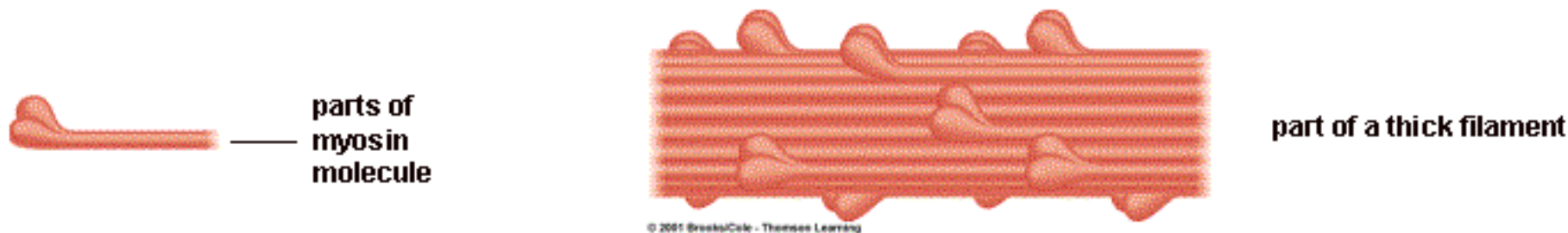
- Like two strands of pearls twisted together
- Pearls are actin
- Other proteins in grooves in filament

Thick filaments

- Composed of myosin
- Each myosin molecule has tail and a double head



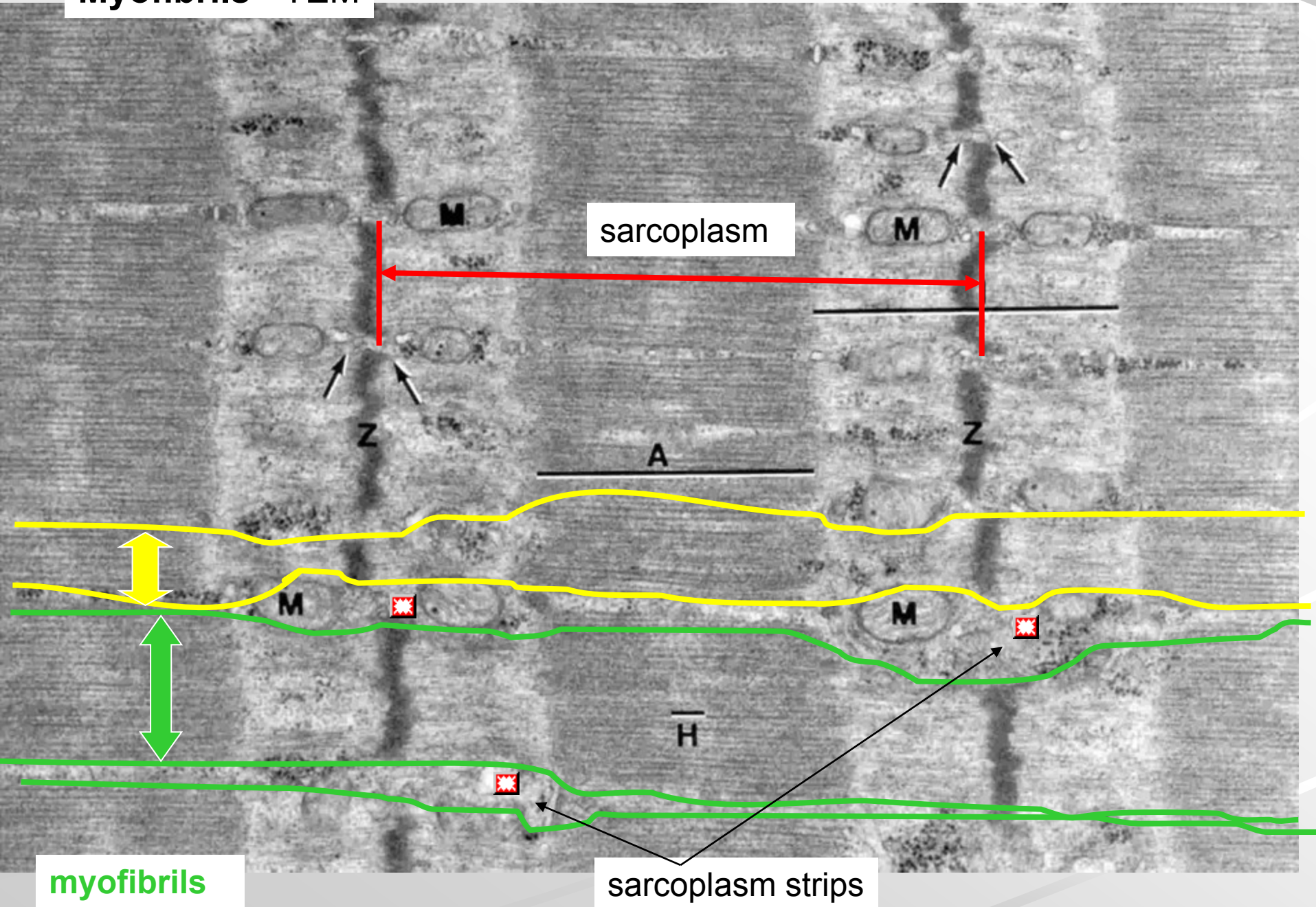
Arrangement of actin molecules in the thin filaments of a sarcomere



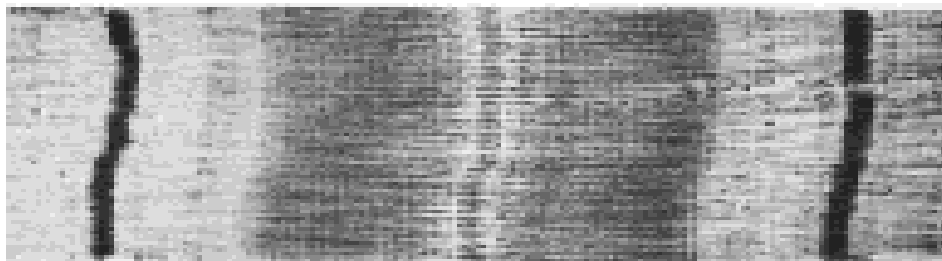
Arrangement of myosin molecules in the thick filaments of a sarcomere



Myofibrils - TEM



Sarcomere

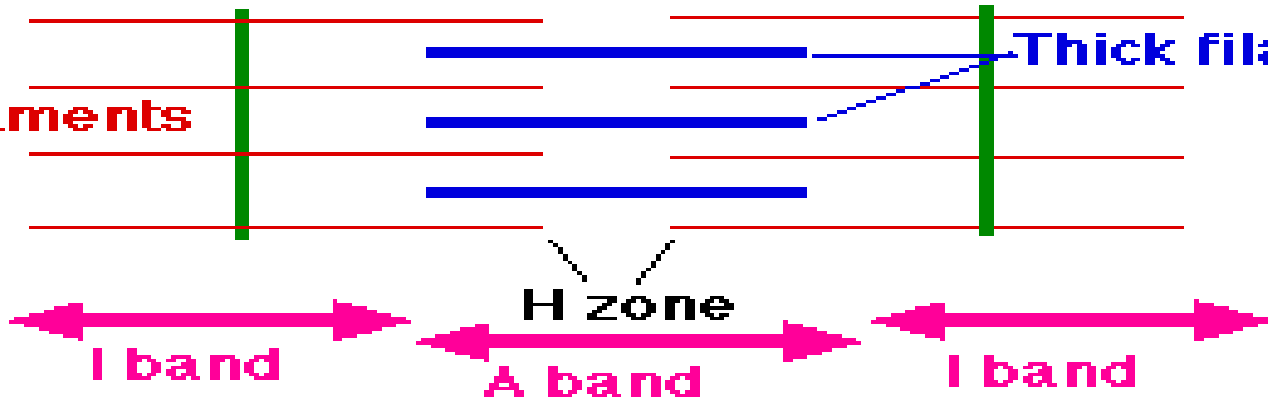


Z line

Z line

Thin filaments

Thick filaments



H zone

I band

A band

I band

A-band

M-line

I-band

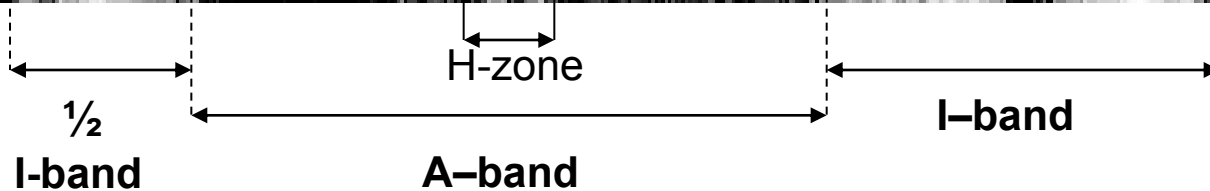
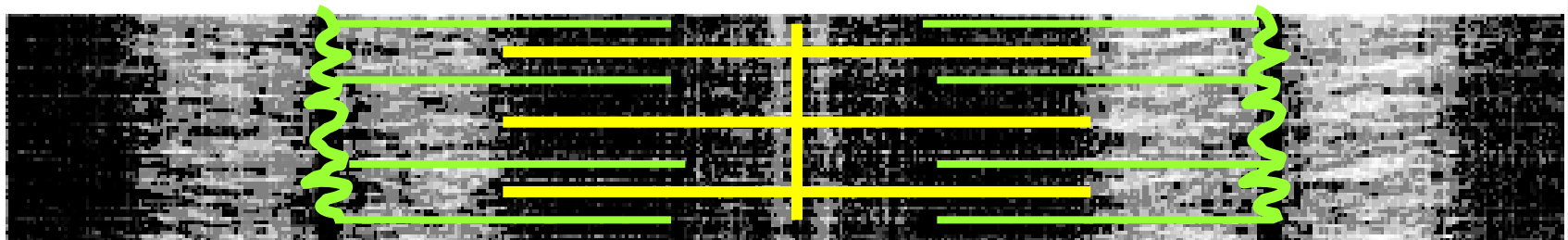
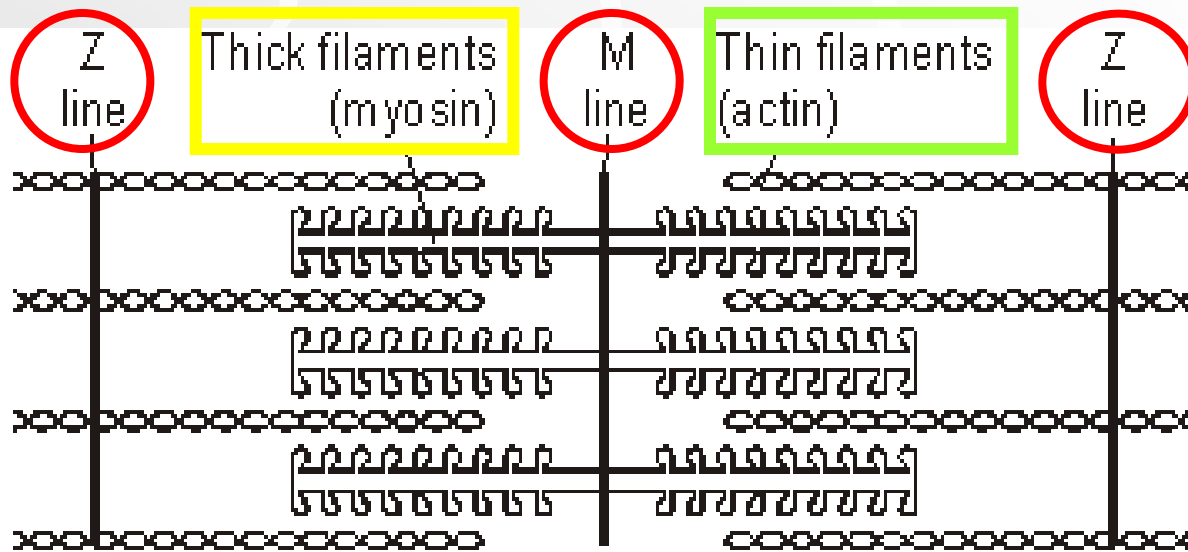


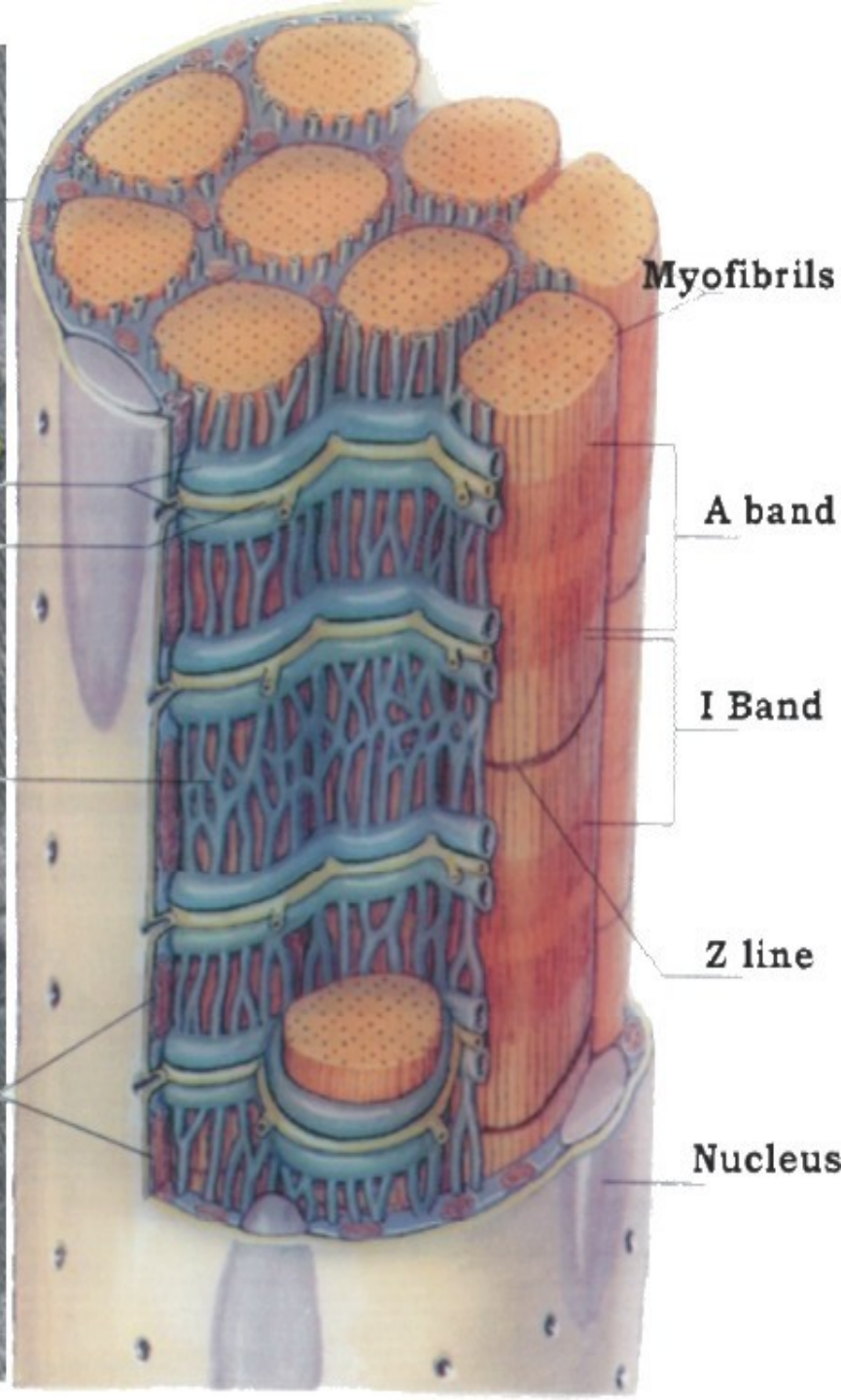
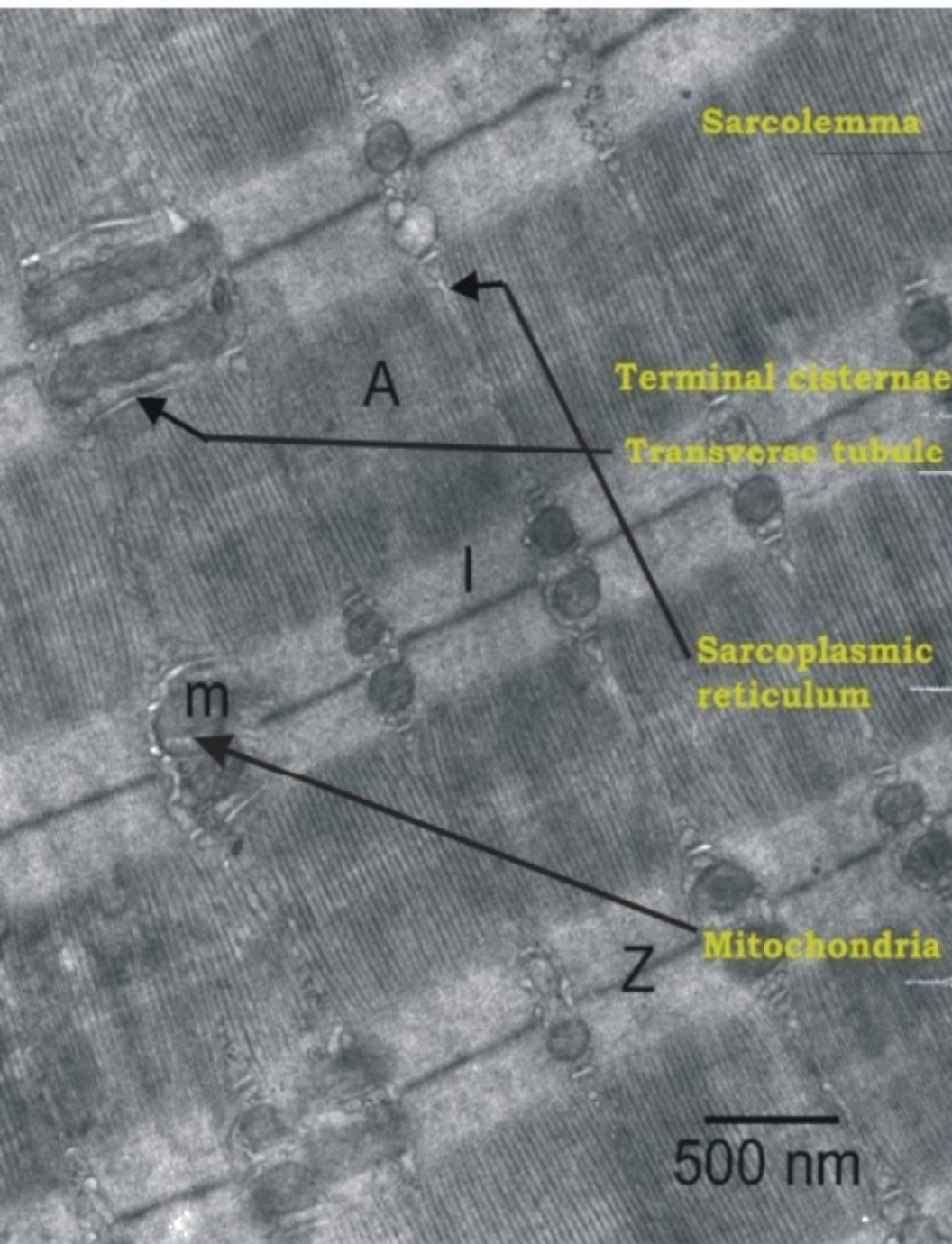
Z-disc

Sarcomere

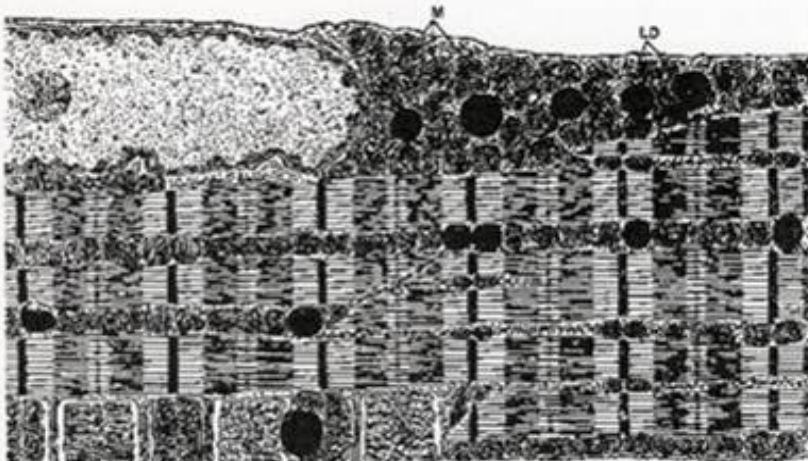
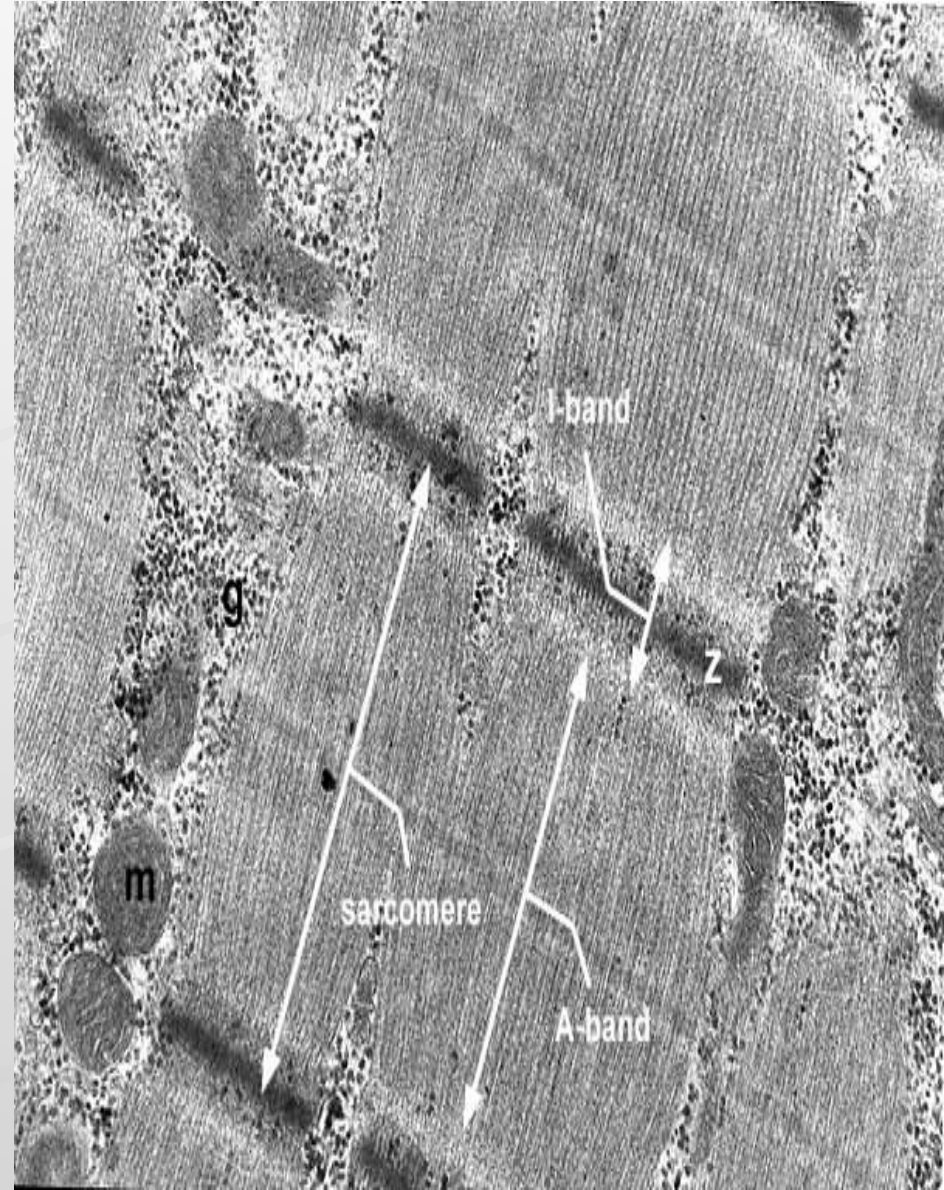
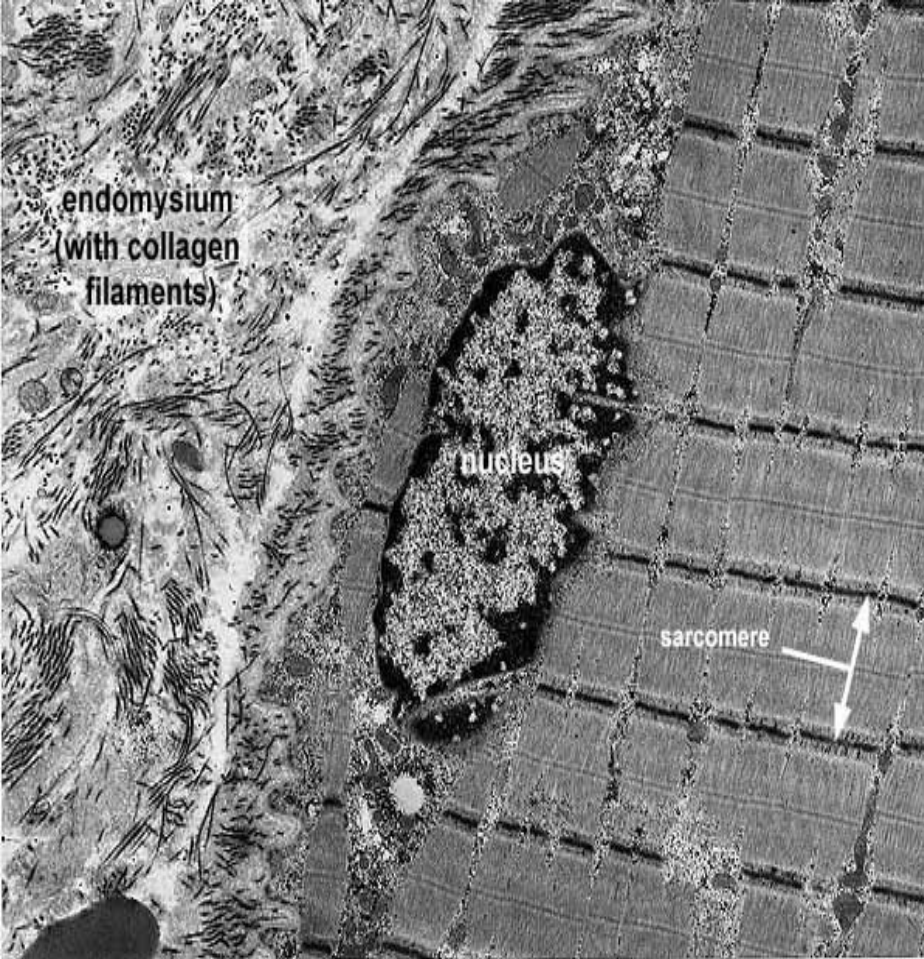
Z-disc

Sarcomere





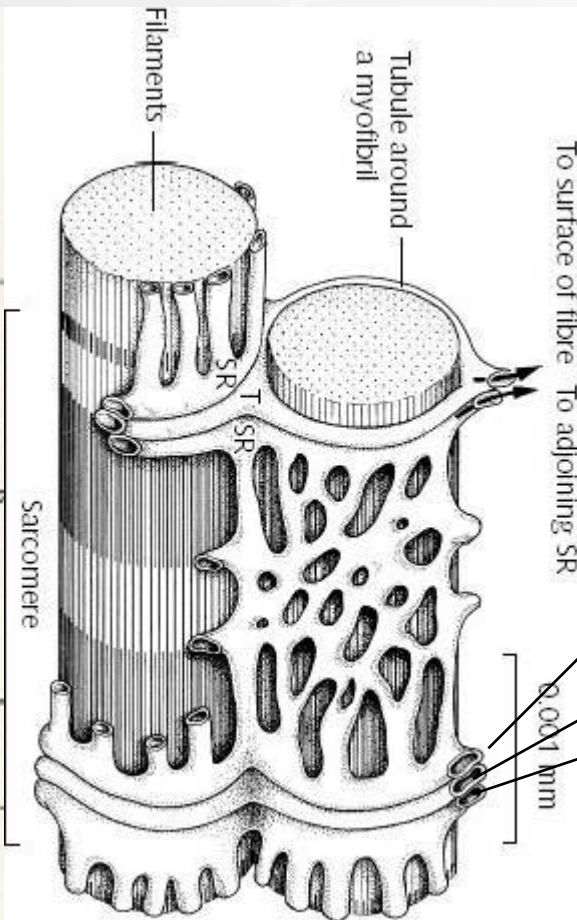
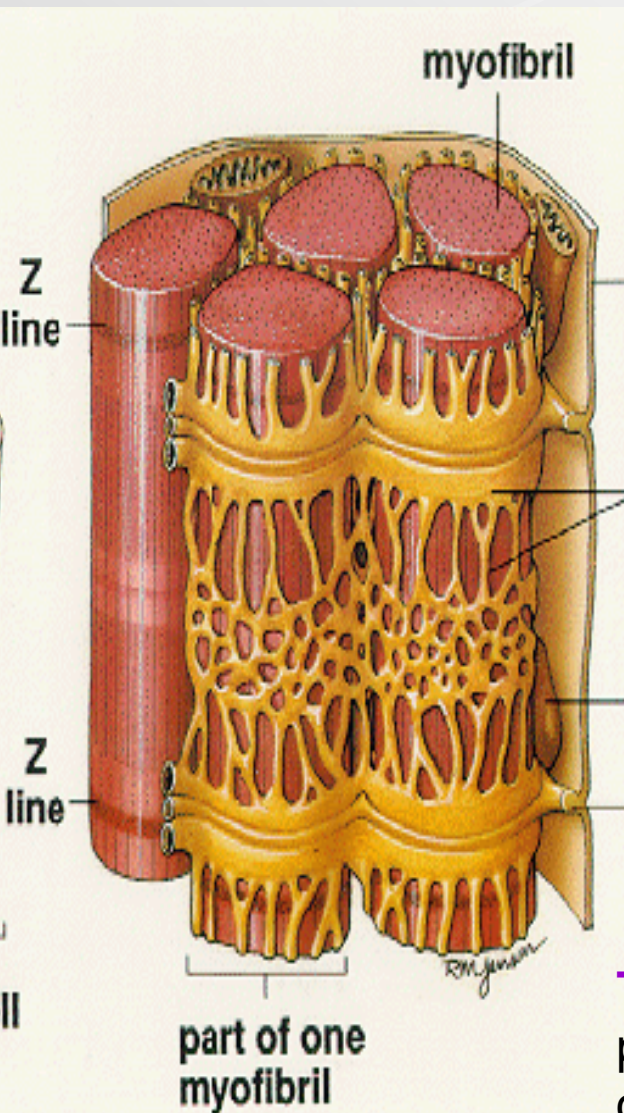
Skeletal muscle in EM



Sarcoplasmic reticulum, t-tubule

Forms **transversal terminal cisternae** and longitudinal tubules.

Function: reservoir of Ca ions



Terminal cisterna
T-tubule
Terminal cisterna } **TRIAD**

T-tubules are invaginations of sarcoplasm and leads action potential to terminal cisternae (it changes permeability of membrane for Ca ions)

To surface of fibre To adjoining SR

0.001 mm

Tubule around
a myofibril

Filaments

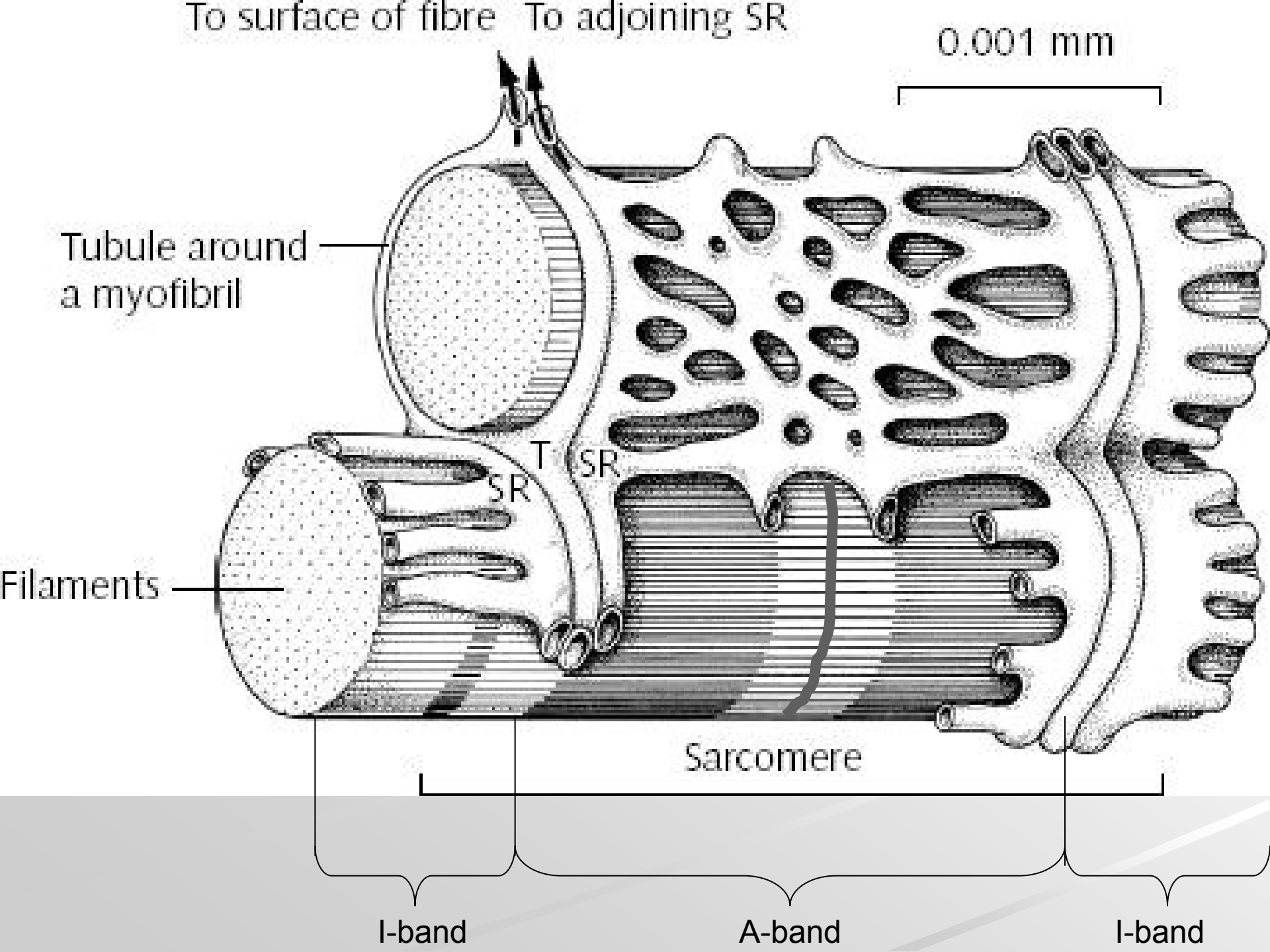
T
SR

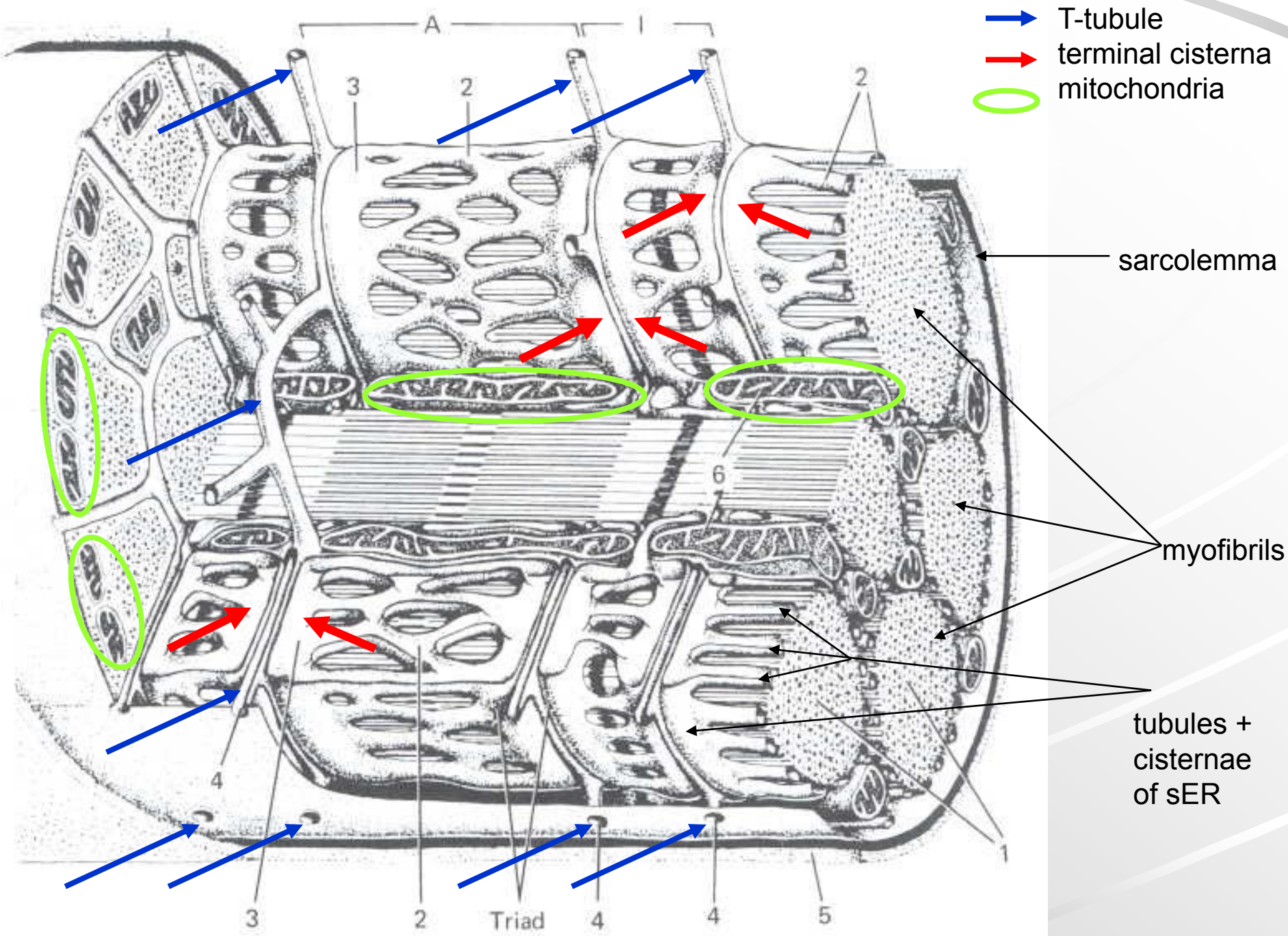
Sarcomere

I-band

A-band

I-band





- T-tubule
- terminal cisterna
- mitochondria

sarcolemma

myofibrils

tubules +
cisternae
of sER

3

2

Triad

4

4

5

6

A

I

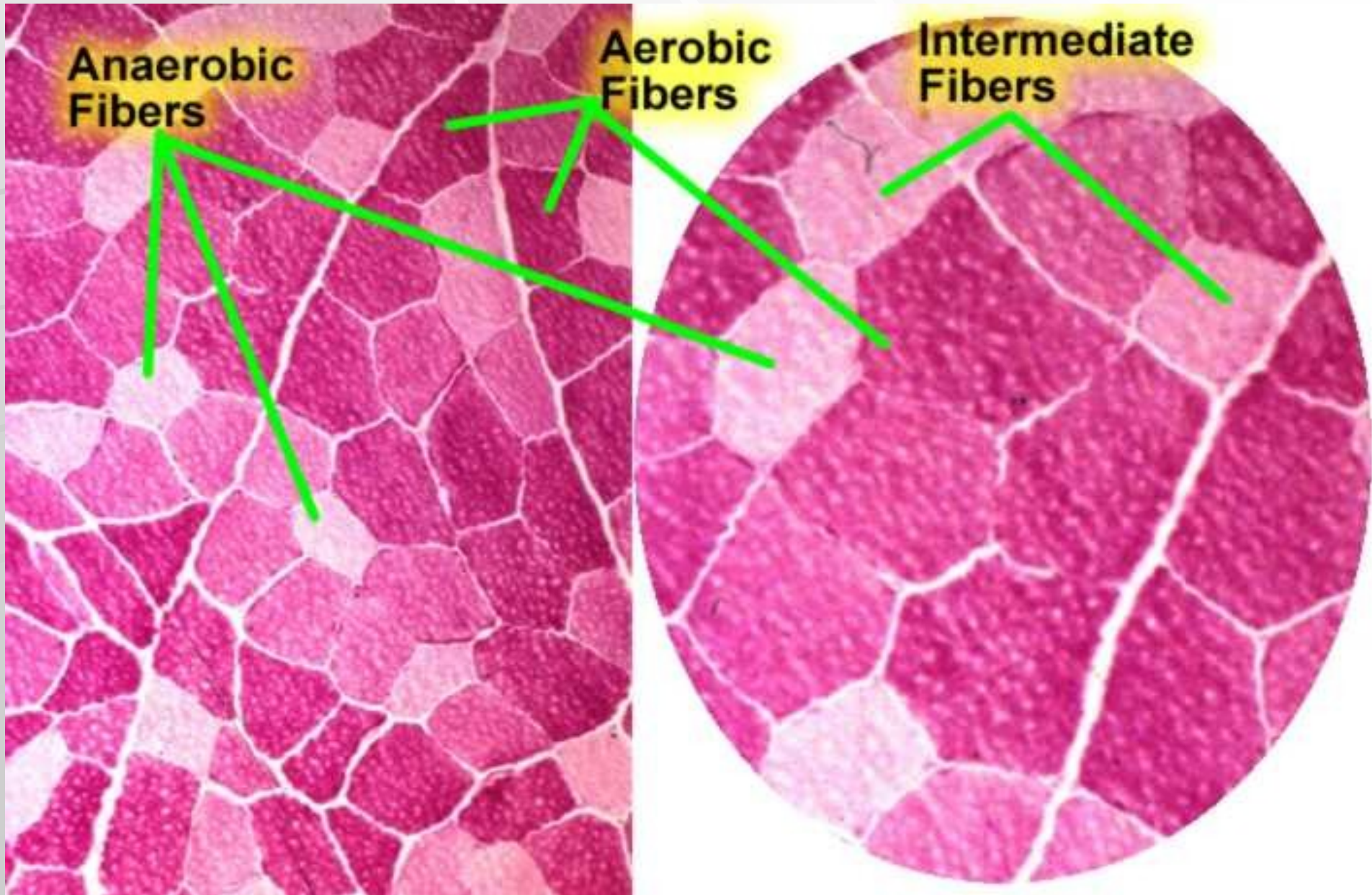
Characteristic of skeletal muscle fiber types

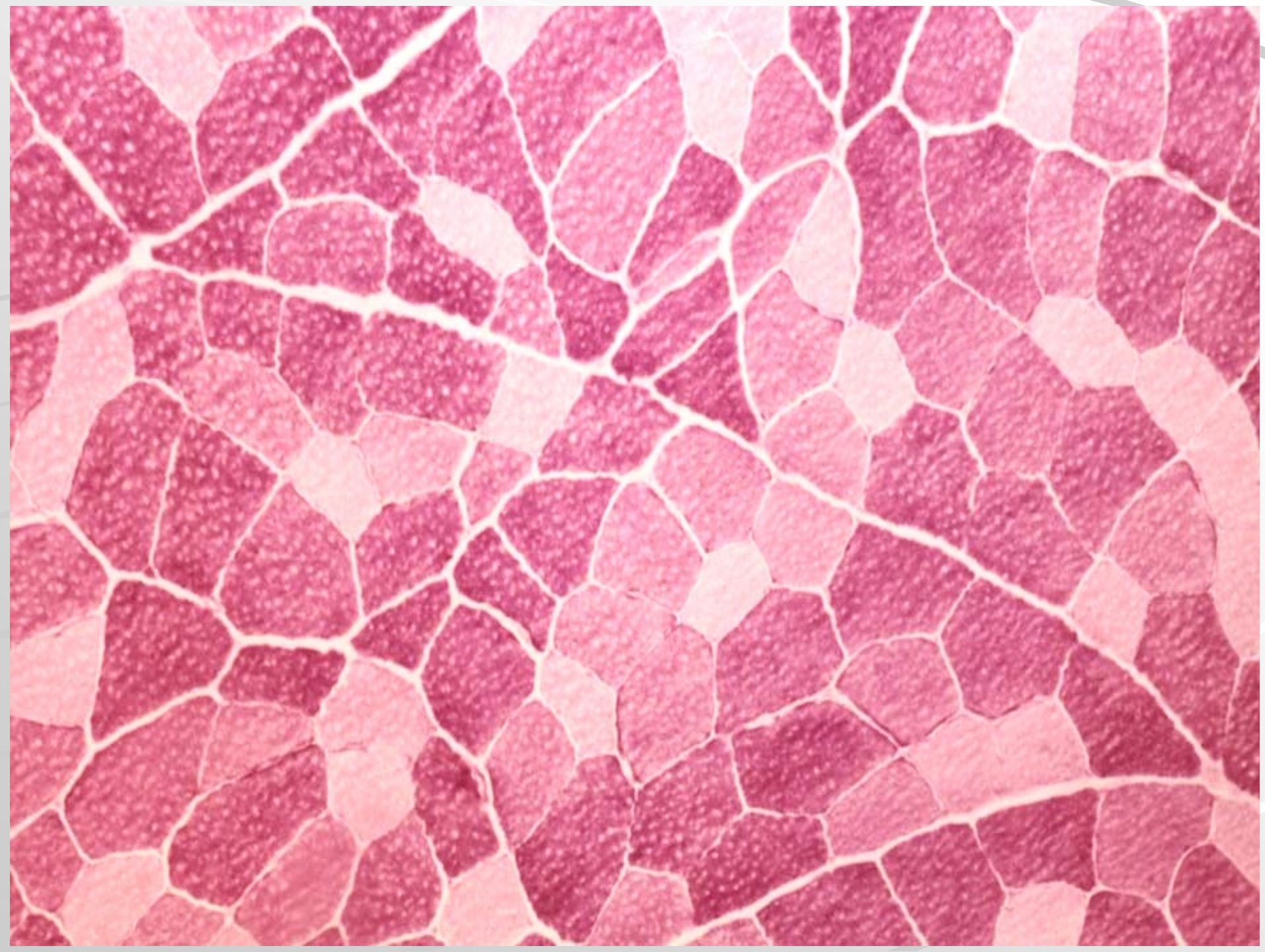
Fibers:	Red	Intermediary	White
Colour	Dark (red)	Dark (red)	Light (white)
Content of myoglobin	High	Intermediate	Low
Number of mitochondria	Many small Mi	Many large Mi	Little small Mi
ATPase activity	Low	High	High
Type of metabolism	Oxidative	Aerobic and anaerobic	Anaerobic
Contraction	Slow	Quick	Quick
Defatigability	Very small	Small	High
Capillary density	High	High	Low

**Anaerobic
Fibers**

**Aerobic
Fibers**

**Intermediate
Fibers**



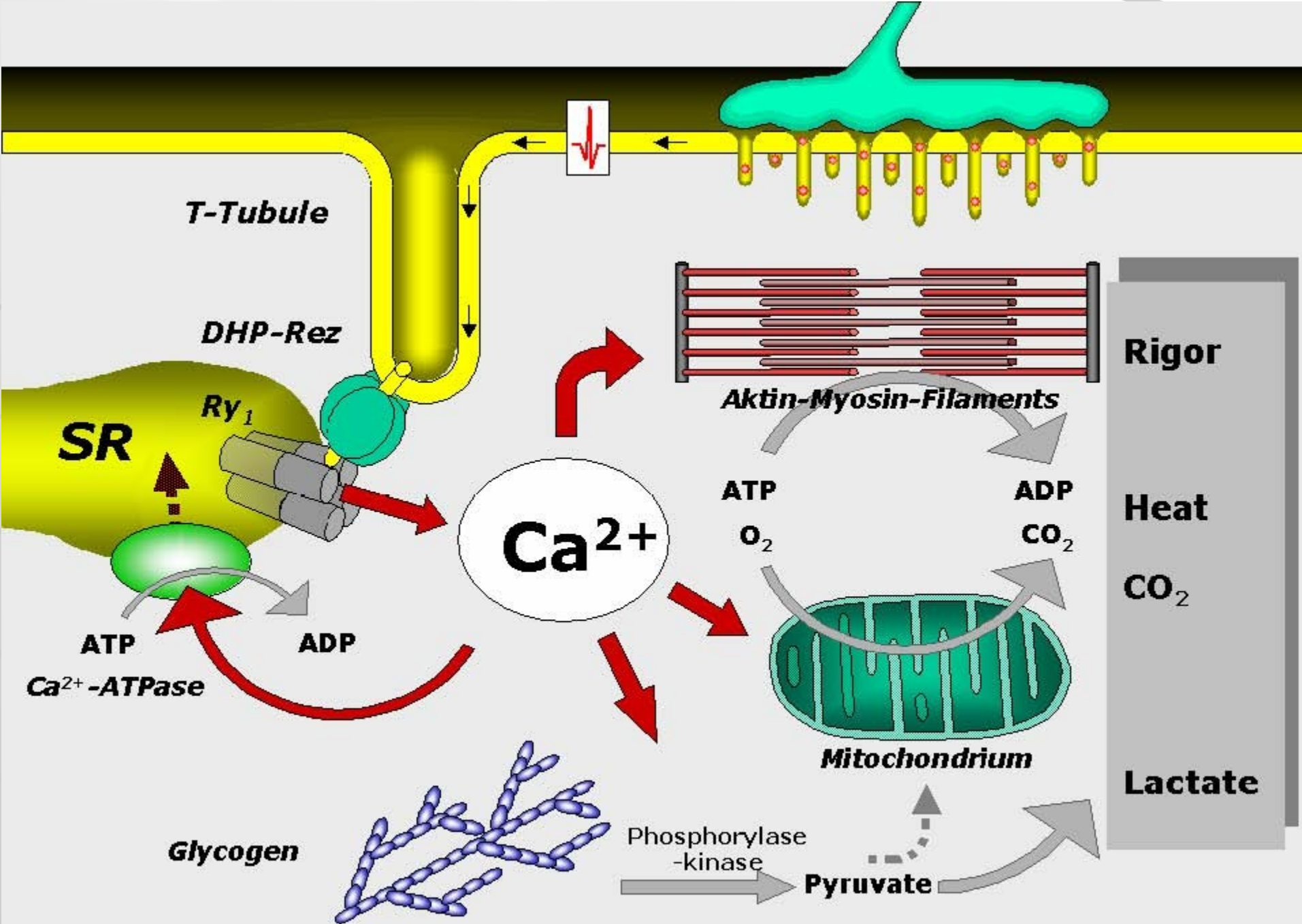


Contraction

- ✦ Propagation of action potential (depolarization) via T-tubule (= invagination of sarcolemma)
- ✦ Change of terminal cisternae permeability – releasing of Ca^+ ions increases their concentration in sarcoplasm
- ✦ Myosin contacts actin and sarcomera shortens by sliding movement – contraction
- ✦ Relaxation: repolarization, decreasing of Ca^{2+} ions concentration, inactivation of binding sites of actin for myosin

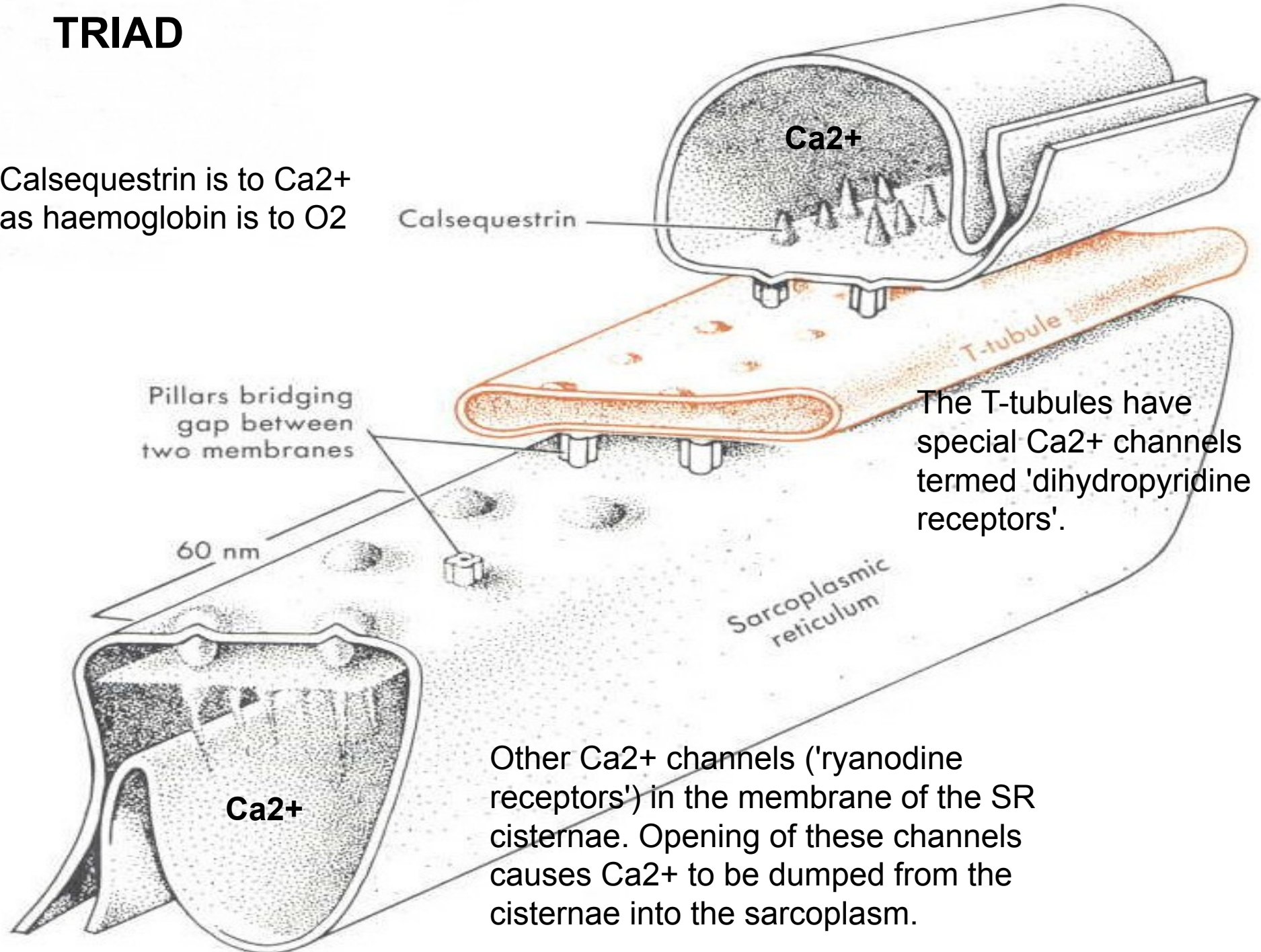


A (ridiculously) simplified model of a sarcomere.



TRIAD

Calsequestrin is to Ca^{2+}
as haemoglobin is to O_2



The T-tubules have special Ca^{2+} channels termed 'dihydropyridine receptors'.

Other Ca^{2+} channels ('ryanodine receptors') in the membrane of the SR cisternae. Opening of these channels causes Ca^{2+} to be dumped from the cisternae into the sarcoplasm.

①

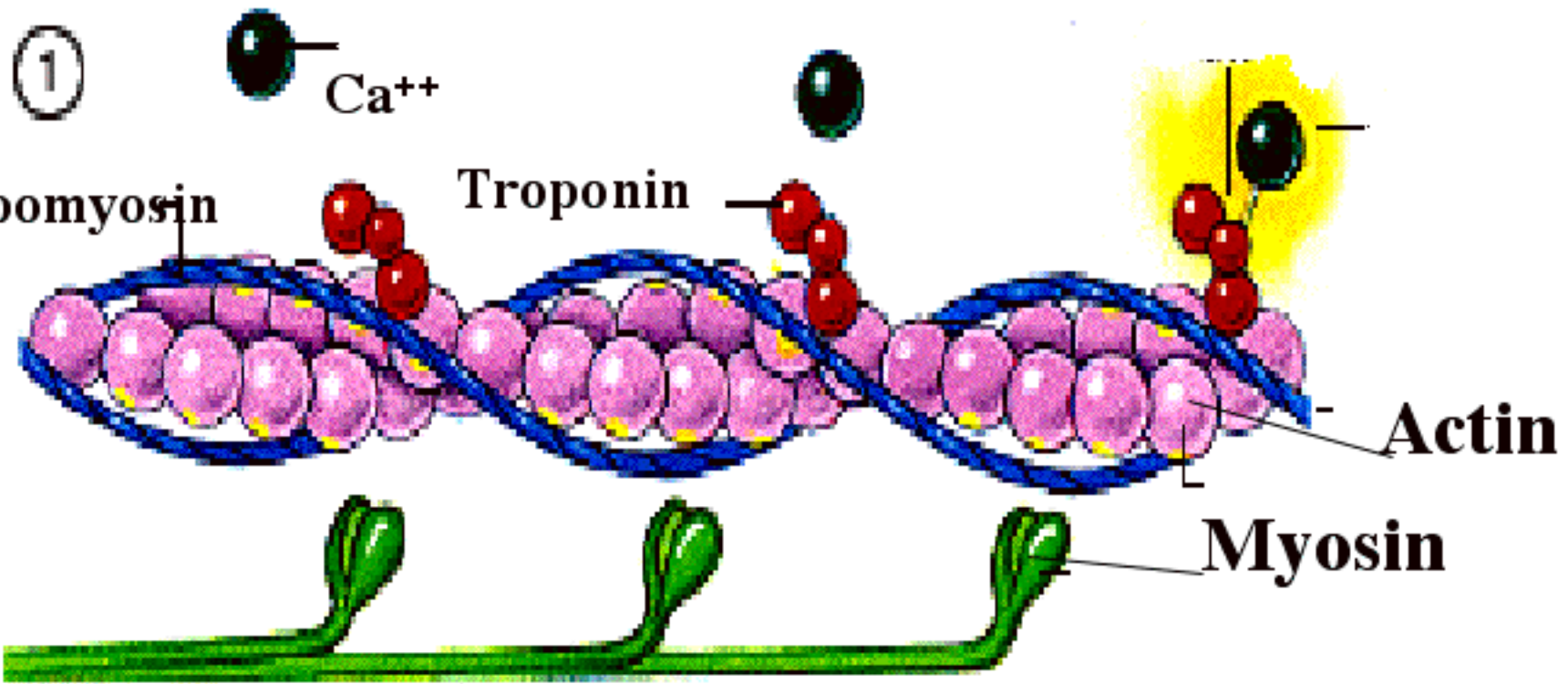
Ca⁺⁺

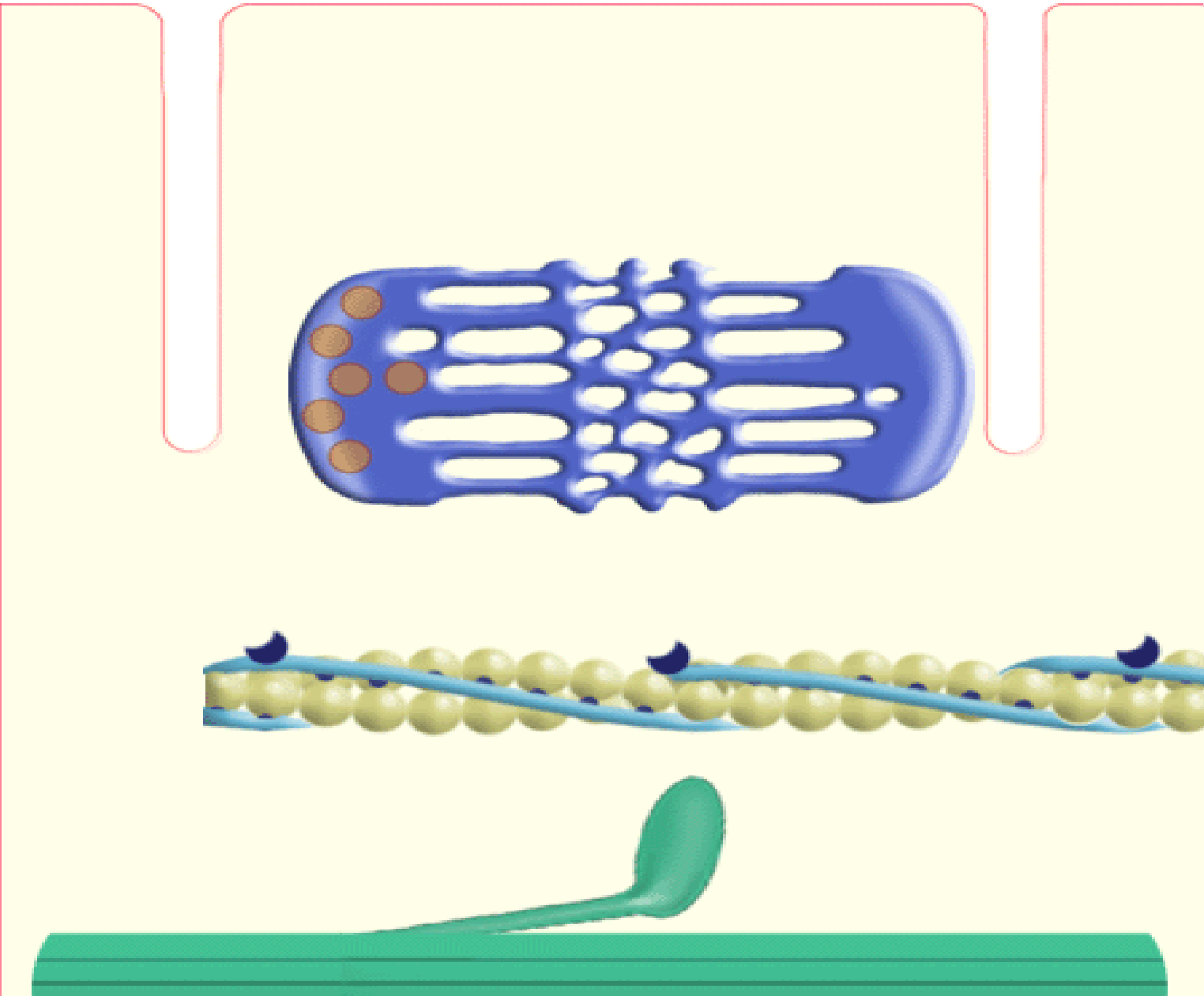
Tropomyosin

Troponin

Actin

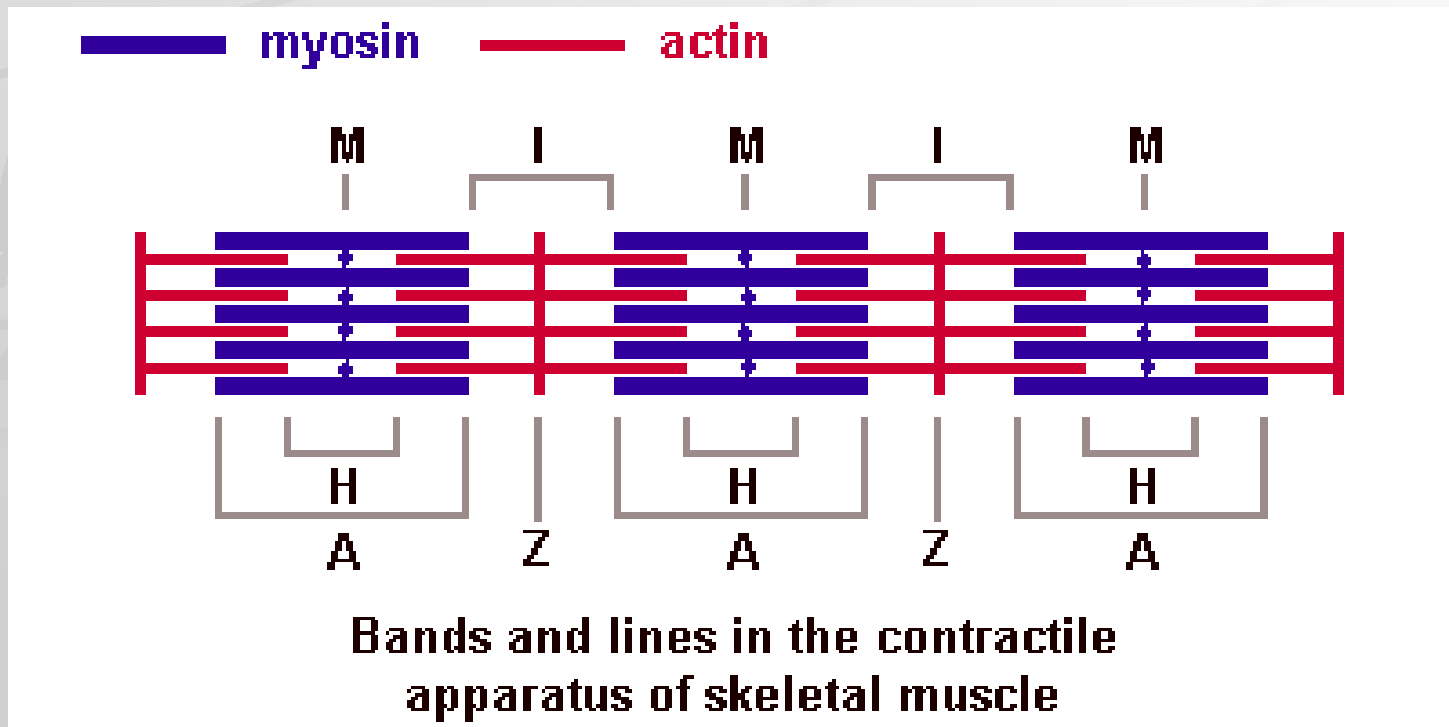
Myosin



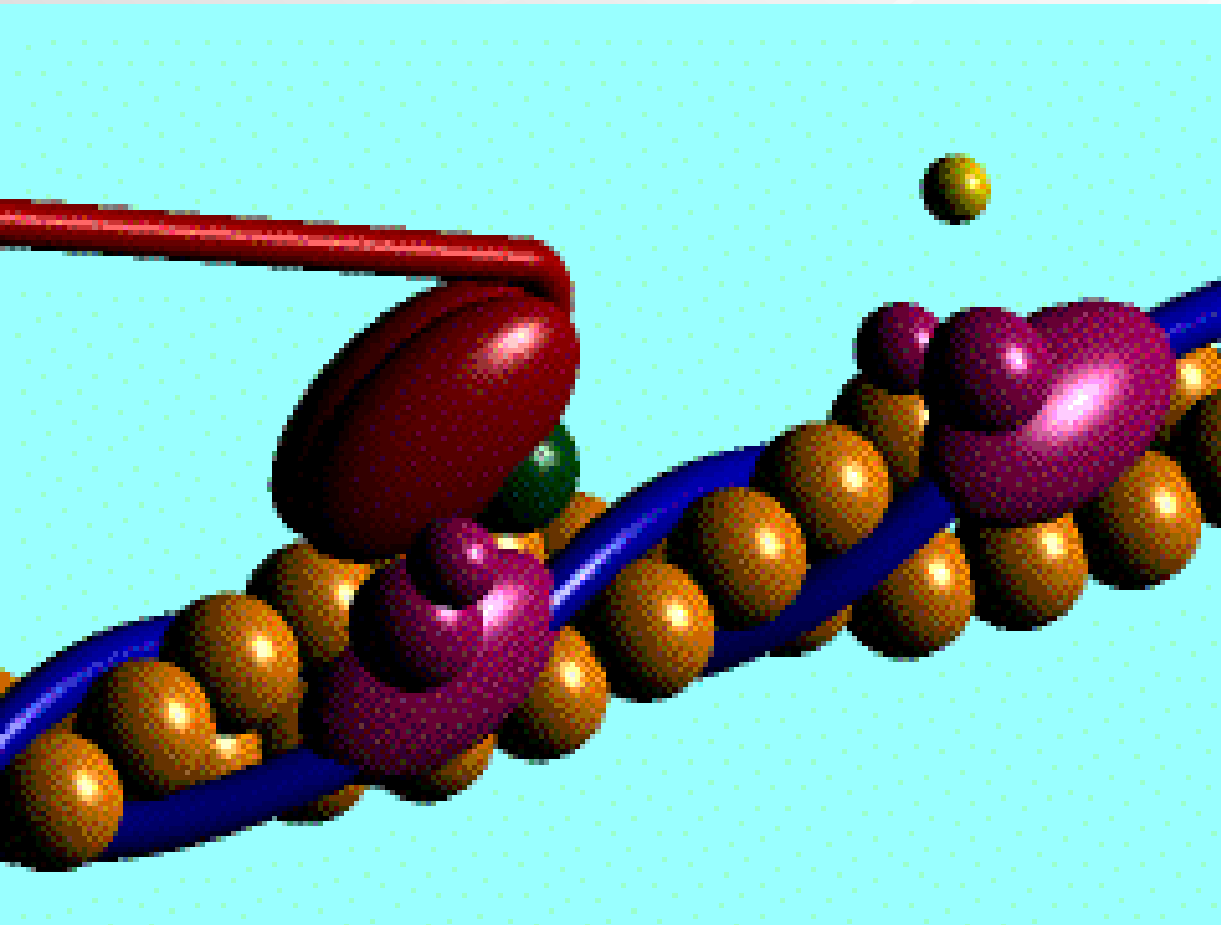


Mechanism of muscle contraction

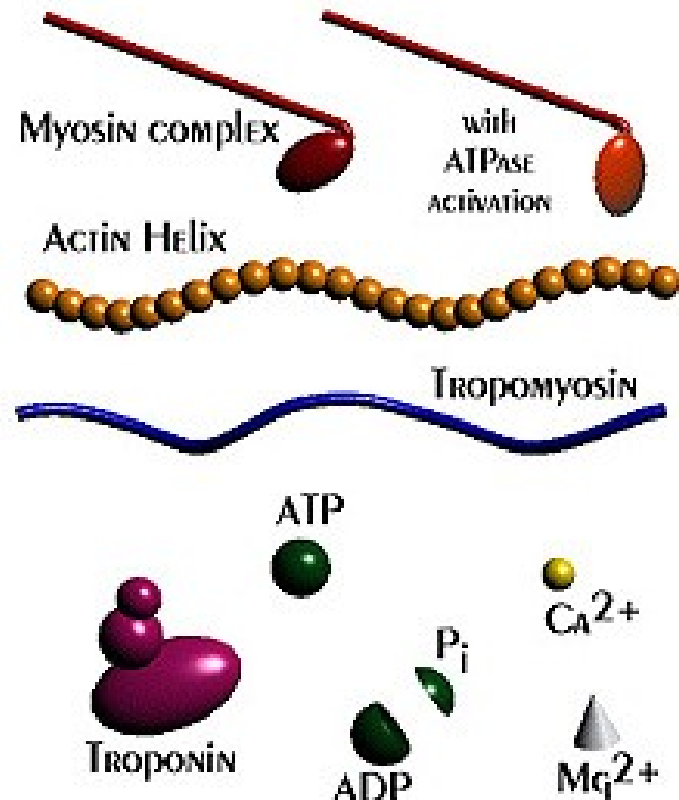
- During contraction the thick /M/ and thin /A/ slide past each other /sliding filament theory/ and the Z-discs are brought near the ends of the thick filaments.



Mechanism of muscle contraction



Players:



Mechanism of contraction: sliding of myofilaments

The Cross Bridge Cycle. (only one myosin head is shown for clarity)

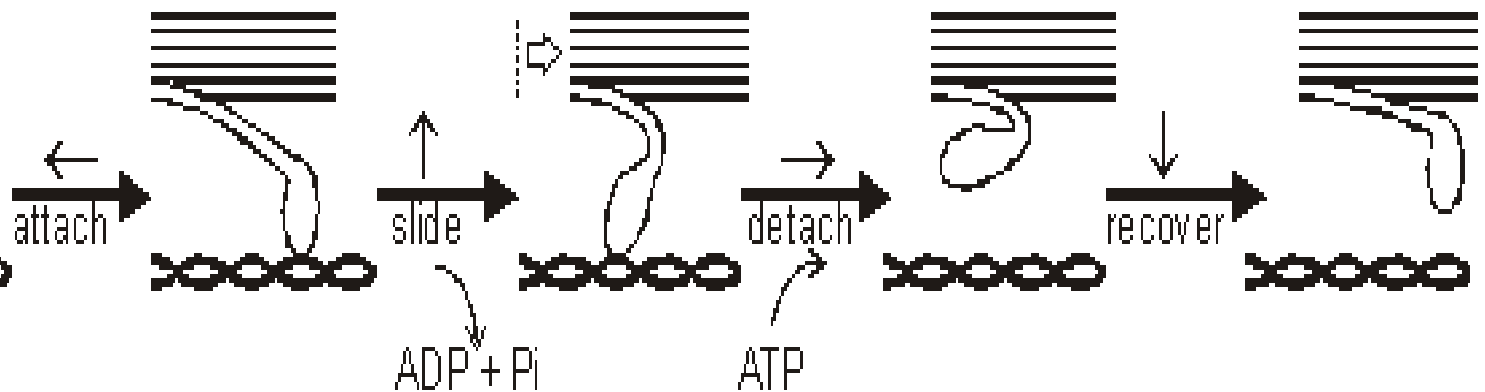
myosin

thick filament:

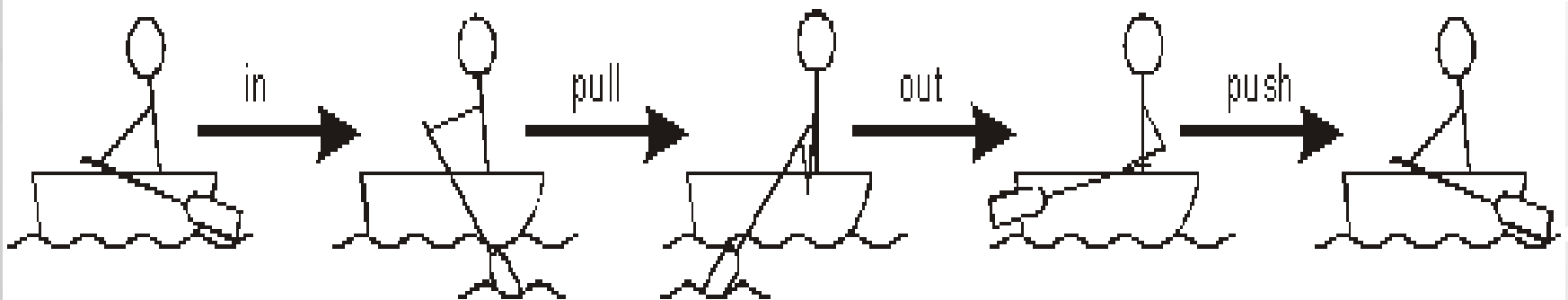
cross
bridge

thin filament

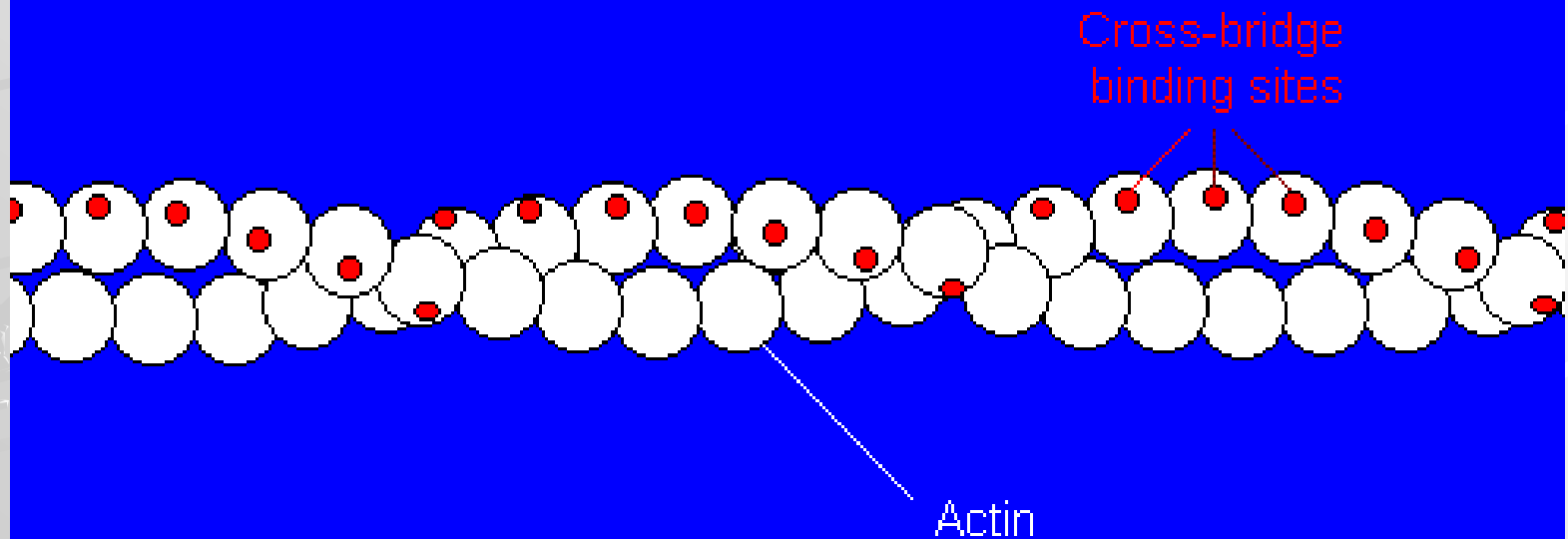
actin



The Rowing Cycle



Cross Bridge Cycle - the Components

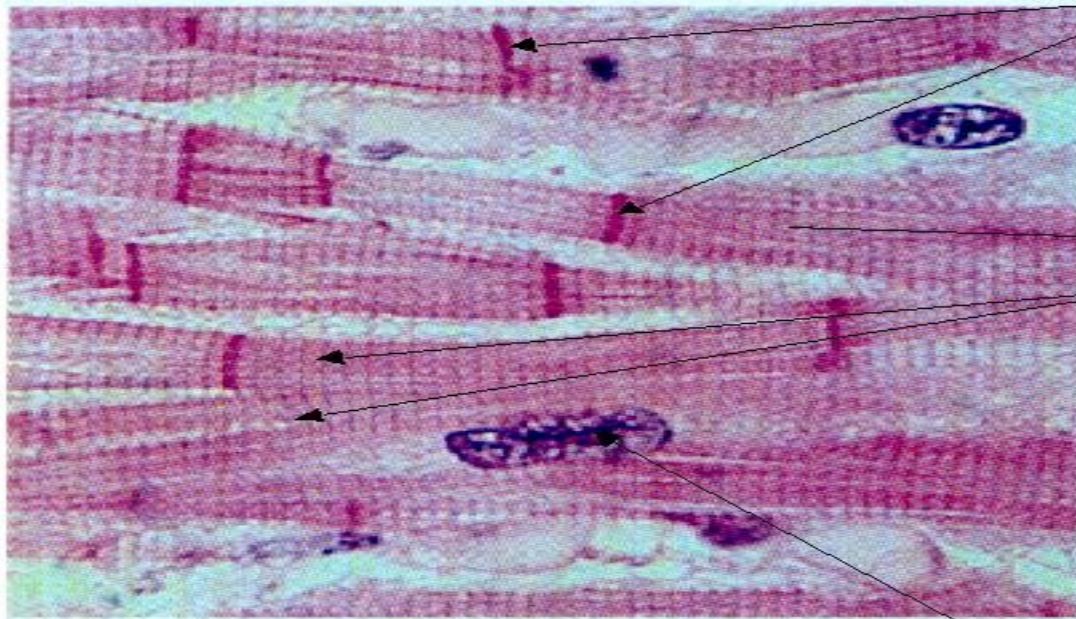


Cardiac muscle - myocardium



- ✚ is made up of long branched fibers, composed of cells – **cardiomyocytes**,
- ✚ cardiomyocytes are cylindrical cells, which can be branched on one or both ends (Y, X shaped cells),
- ✚ Sarkoplasm: 1 nucleus in the center of cell, striated myofibrils, numerous mitochondria,
- ✚ cells are attached to one another by end-to-end junctions – intercalated discs.

Cardiac Muscle Characteristics



Intercalated disks

Cardiac muscle cells are faintly striated, branching cells, which connect by means of intercalated disks to form a functional network.



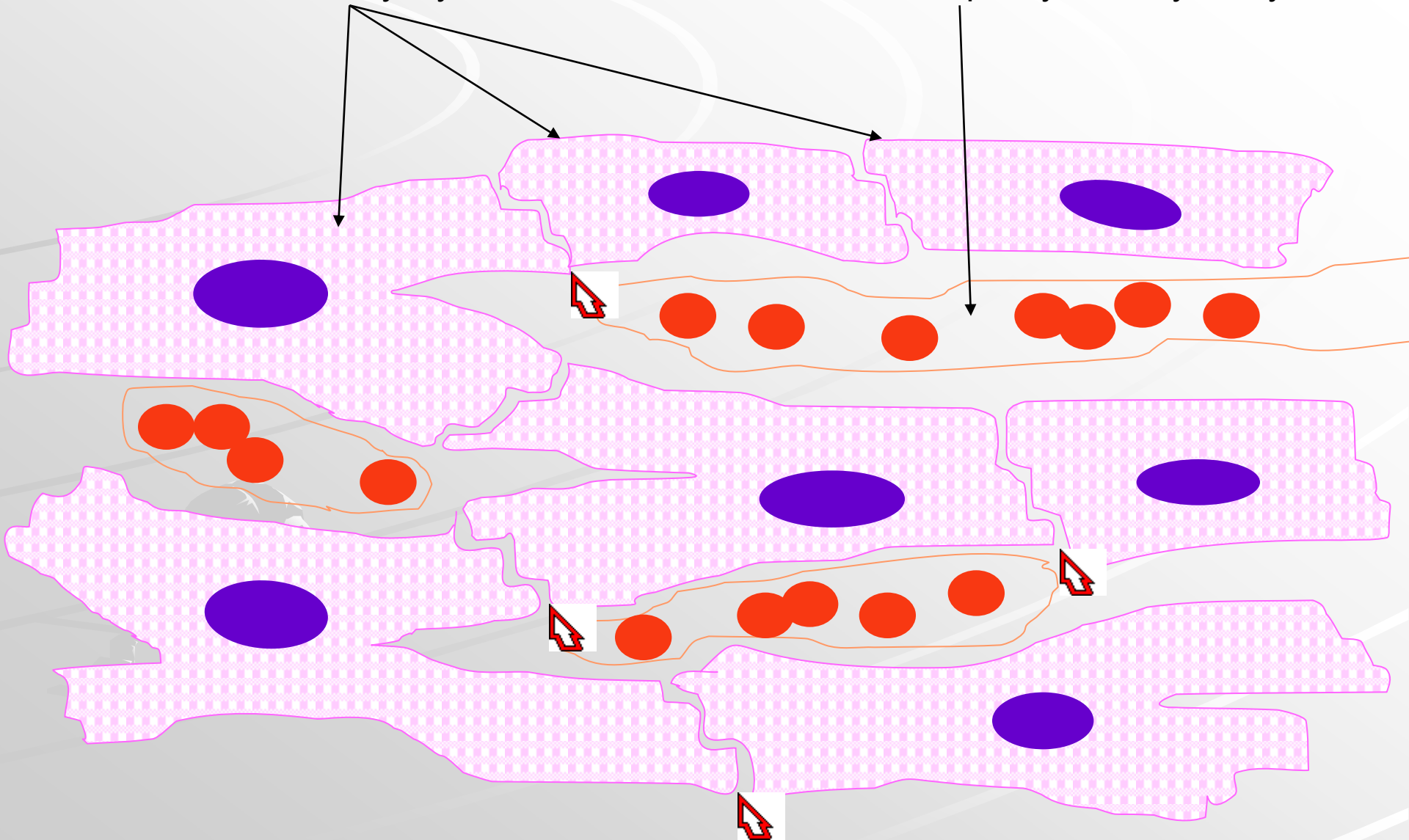
nucleus

Branched, mono-nucleated cells

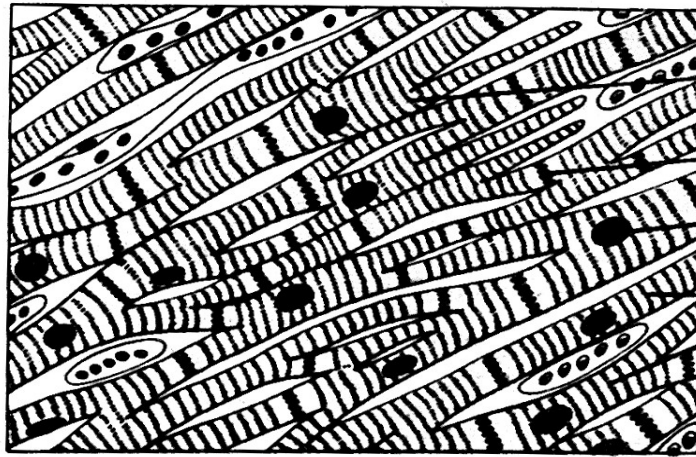
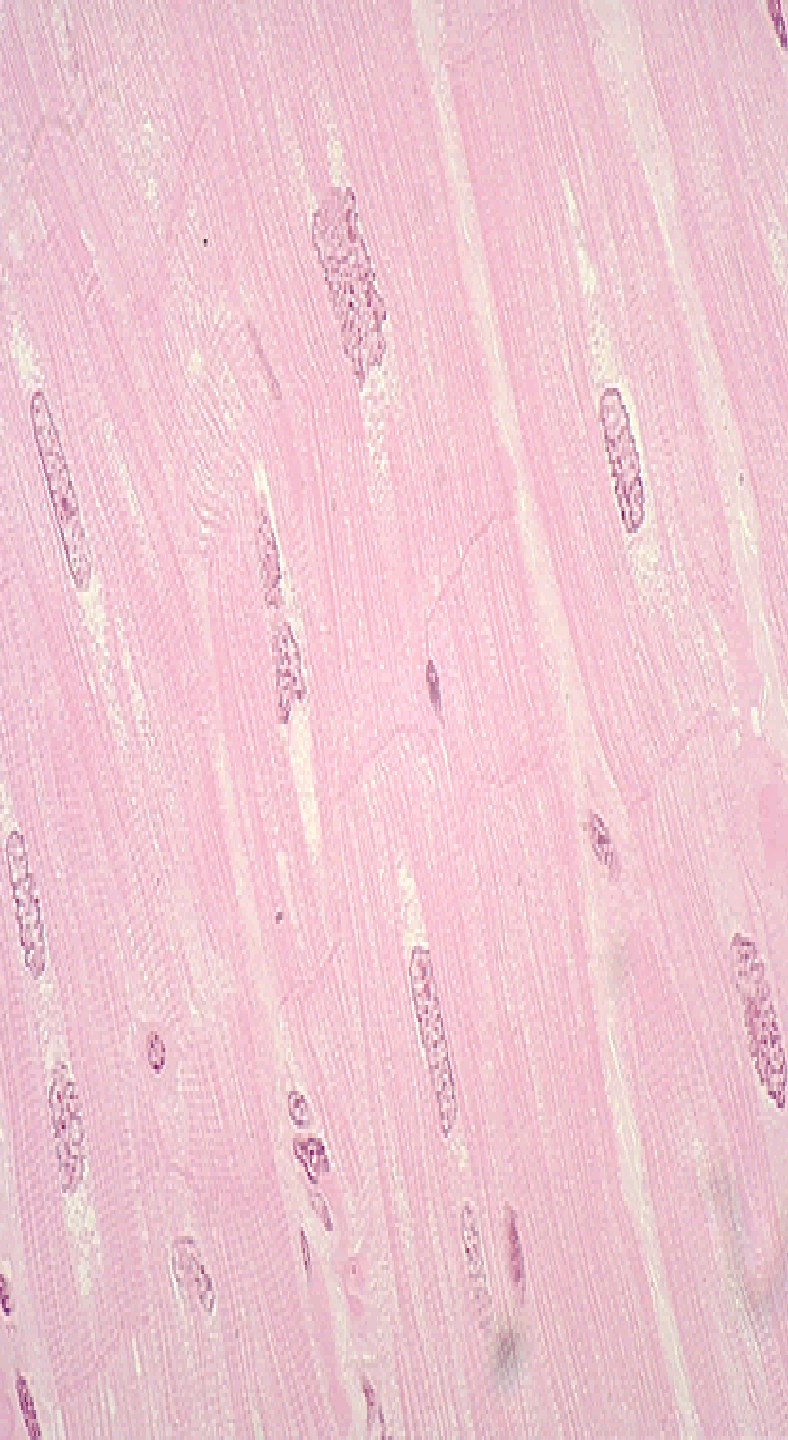
The action potential travels through all cells connected together in the **syncytium** causing them to function as a unit.

chains of cardiomyocytes

blood capillary with erythrocytes



Intercalated disc

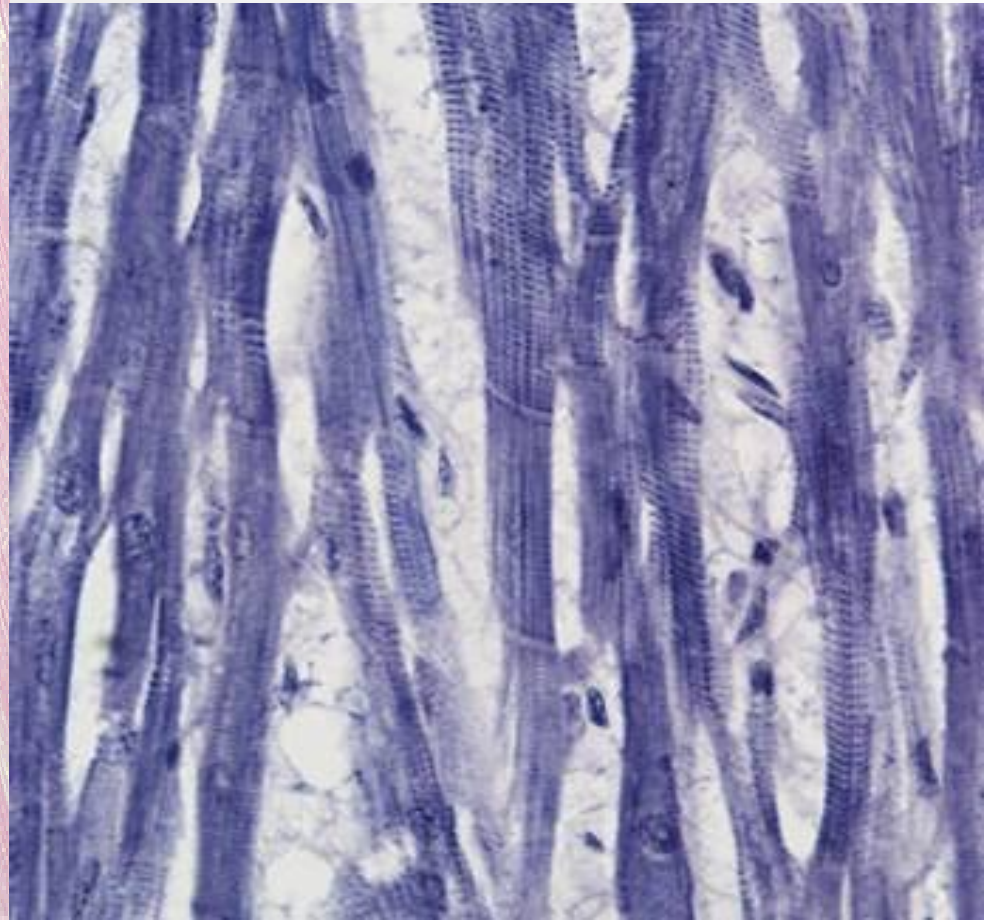


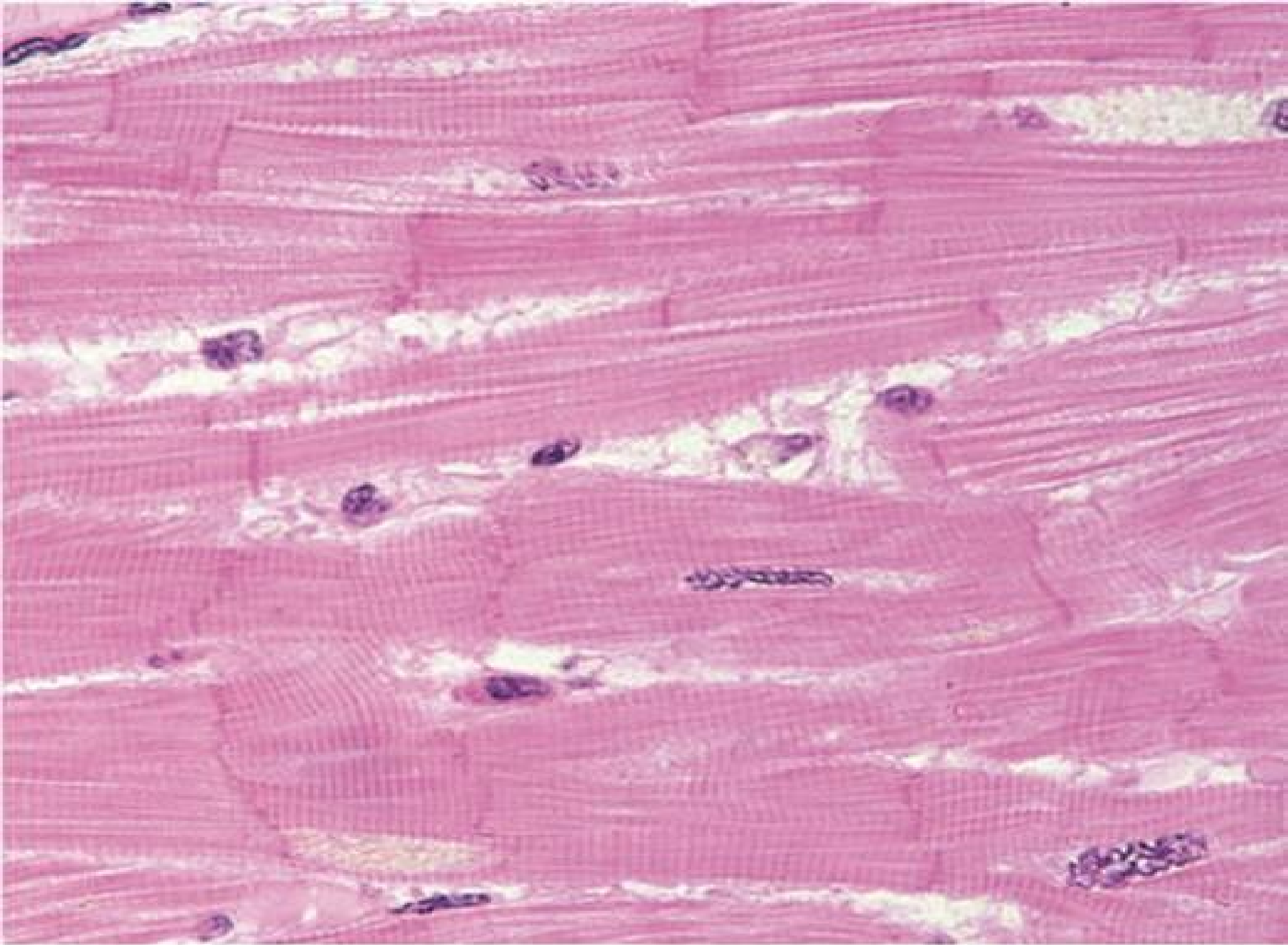
Muscle cell

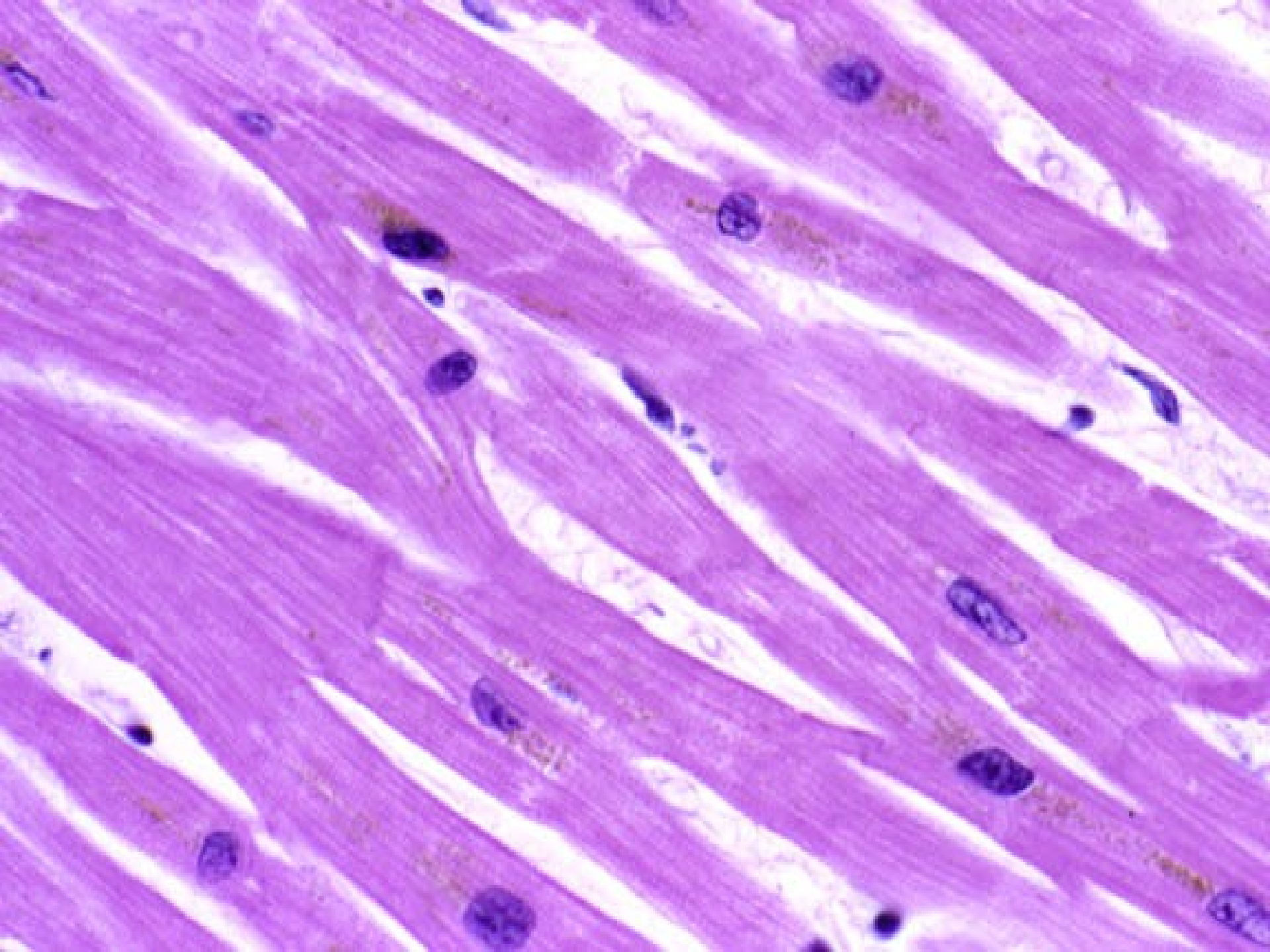
Intercalated disc

Nucleus

Striations



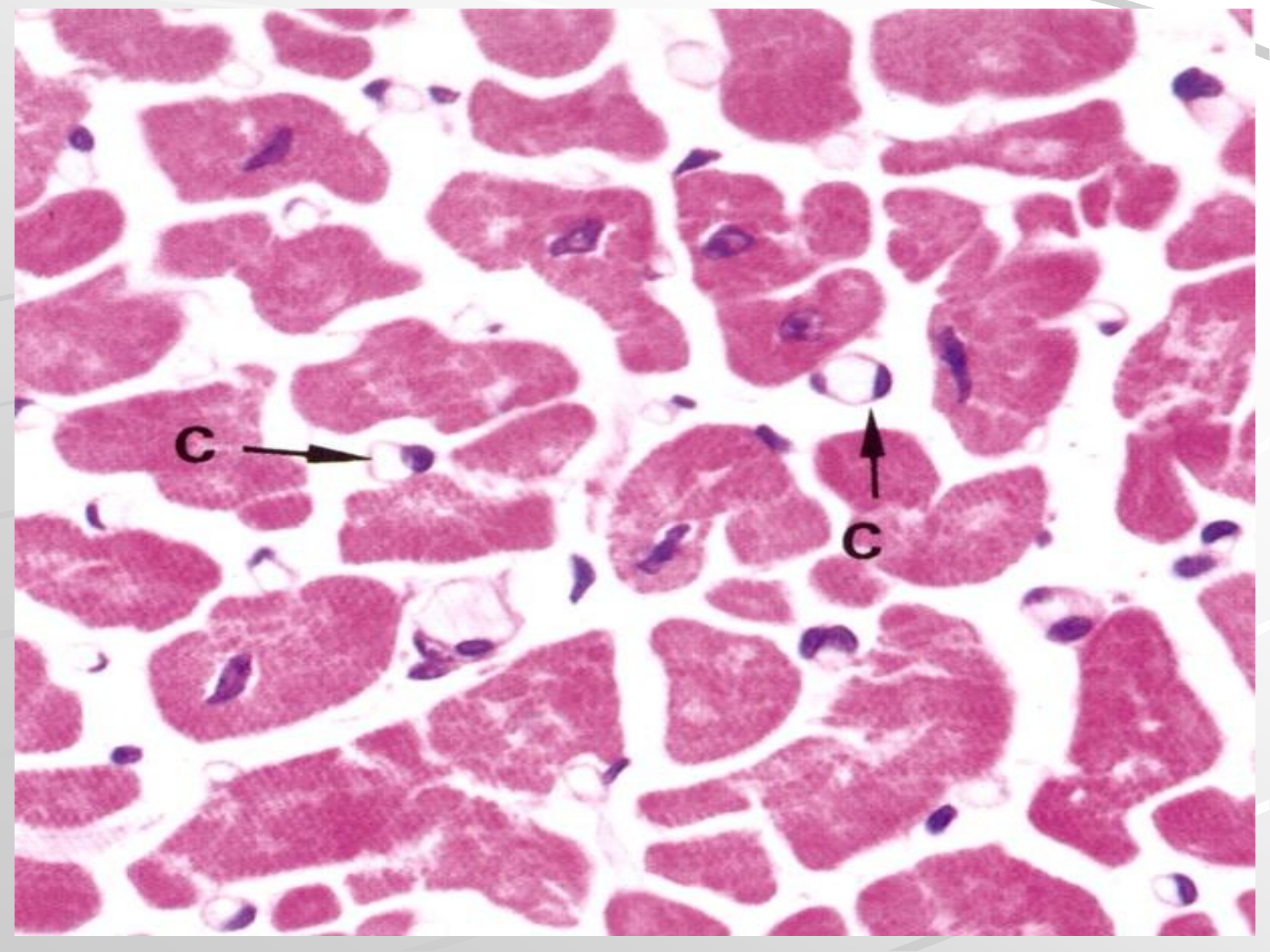




C



C



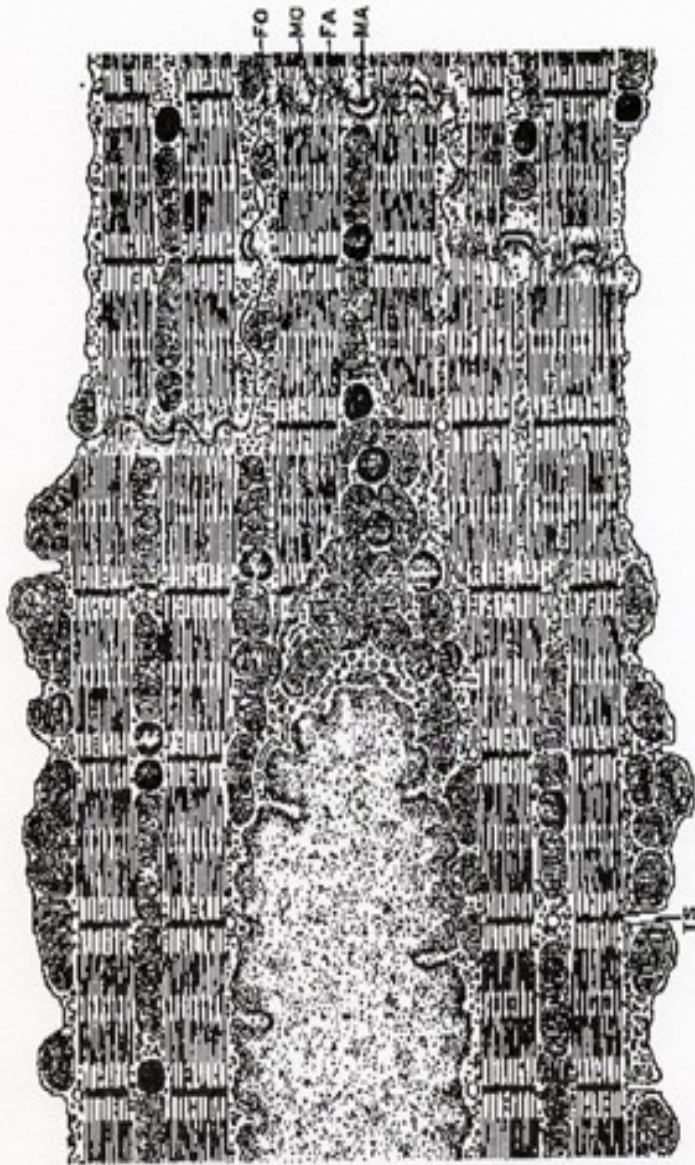
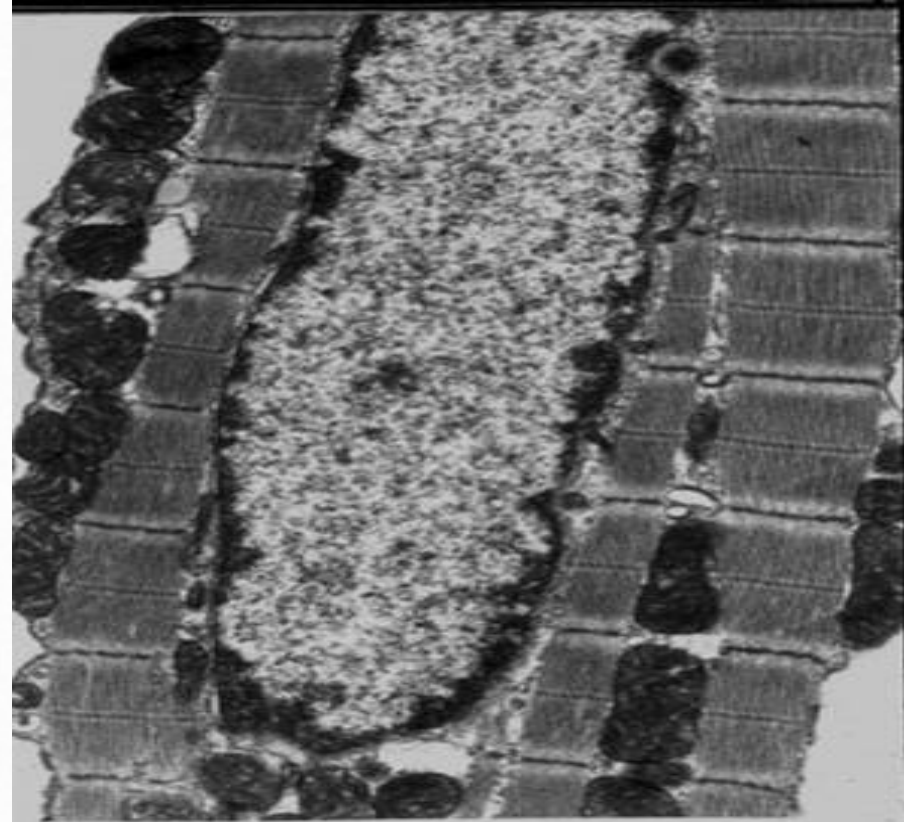


FIG. 10-10 CARDIAC MUSCLE

Diagram and electron micrograph
of a part of cardiomyocyte



DIFFERENCES BETWEEN CARDIAC AND SKELETAL MUSCLE TISSUES

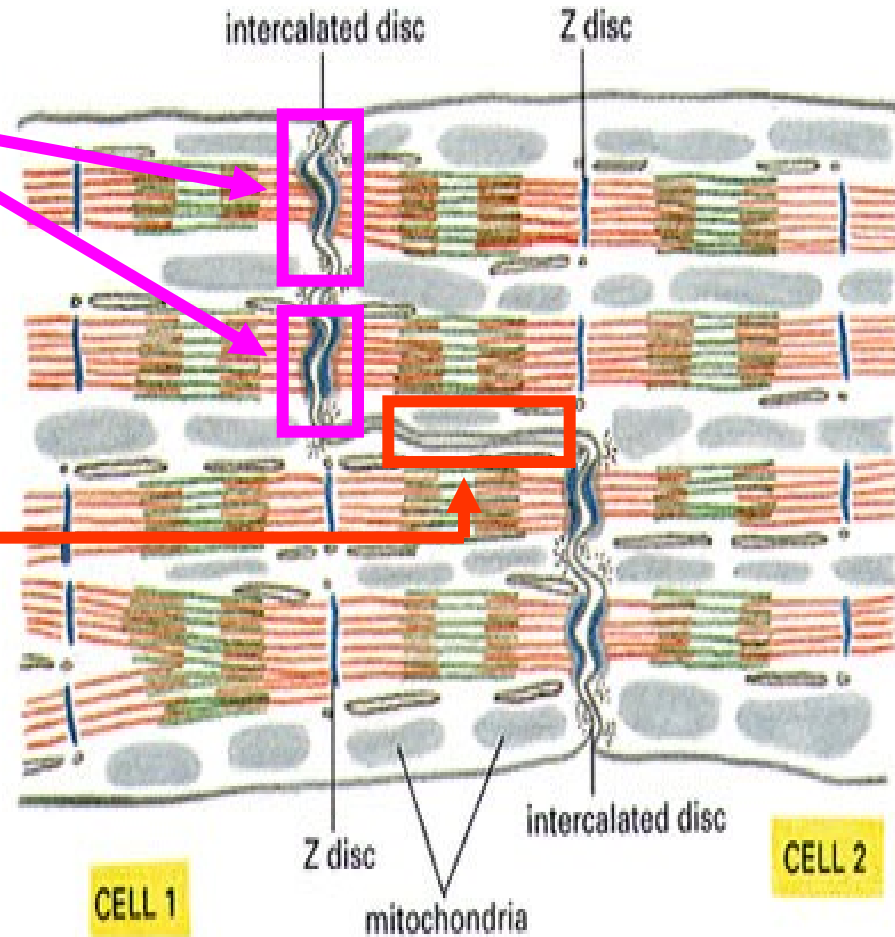
- ✦ there are no triads, but **diads**: 1 t-tubule + 1 cisterna
- ✦ **t-tubules** encircle the sarcomeres **at the Z lines** rather than at the zone of overlap.
- ✦ sarcoplasmic reticulum via its tubules contact sarcolemma as well as the t-tubules
- ✦ cardiac muscle cells are totally dependent on aerobic metabolism to obtain the energy needed to continue contracting. The sarcoplasm thus contains large numbers of **mitochondria** and abundant reserves of **myoglobin** (to store oxygen). Energy reserves are maintained in the form of **glycogen** and **lipid inclusions**.

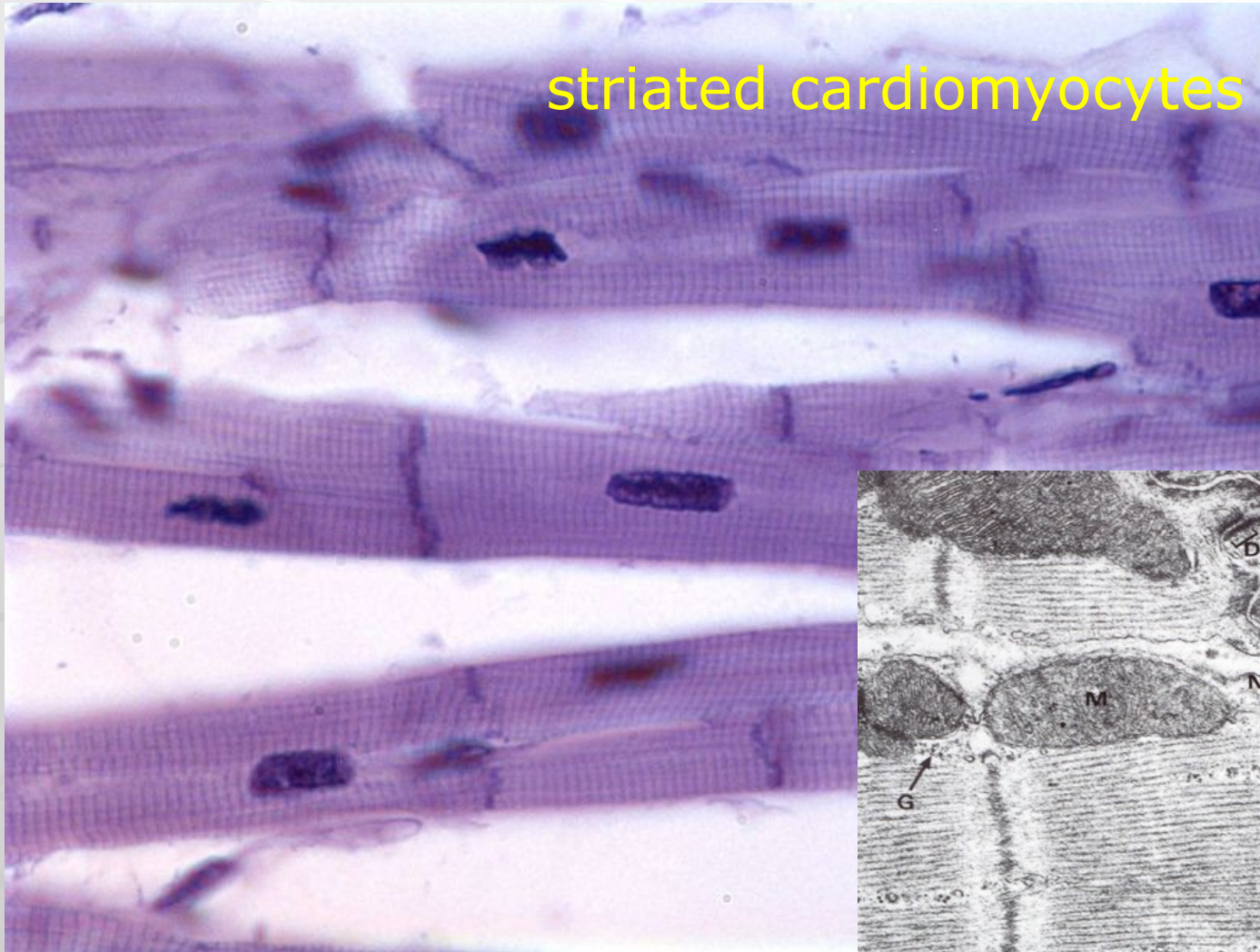
Intercalated disc

„scalariform“ shape of cell ends

fasciae adherentes (*adhesion of cells*)

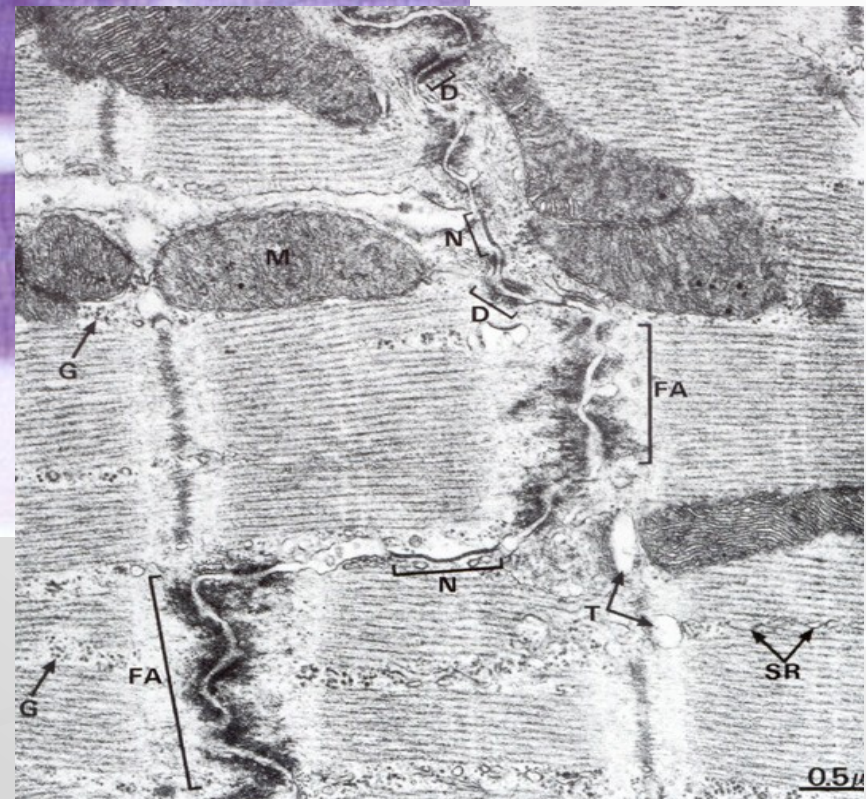
Nexus
(quick intercellular communication – transport of ions, electric impulses, informations)

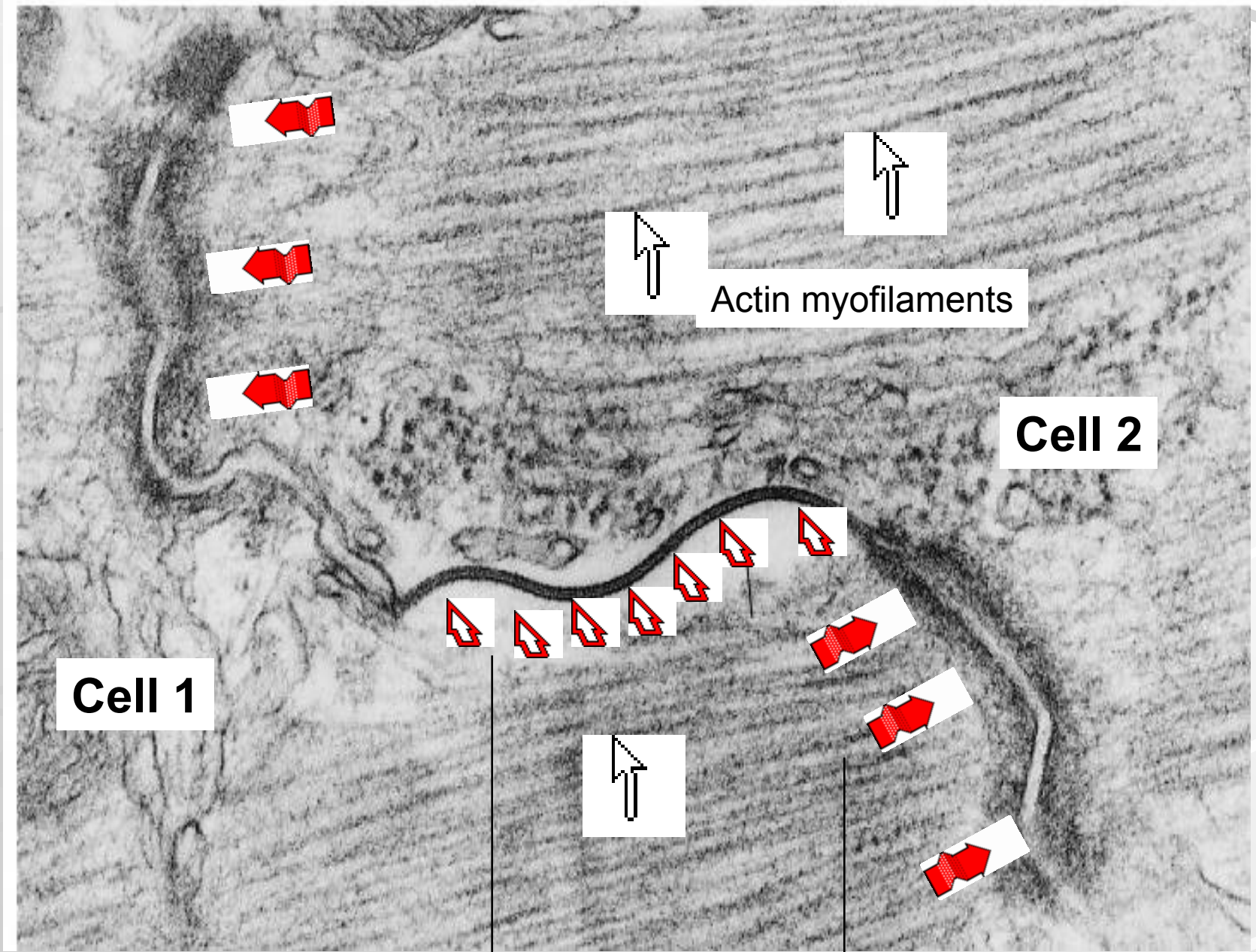




striated cardiomyocytes

intercalated discs –
desmosomes, fasciae adherentes,
nexus





Actin myofilaments

Cell 2

Cell 1

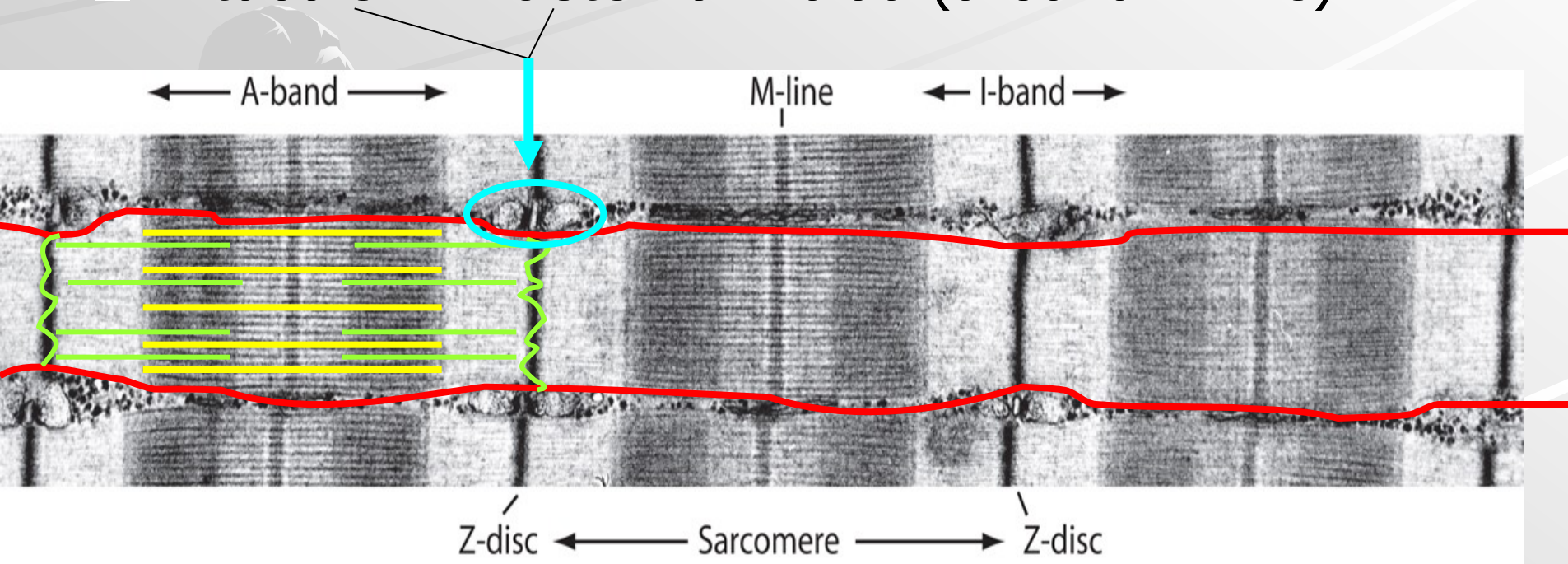
Intercalated disc:

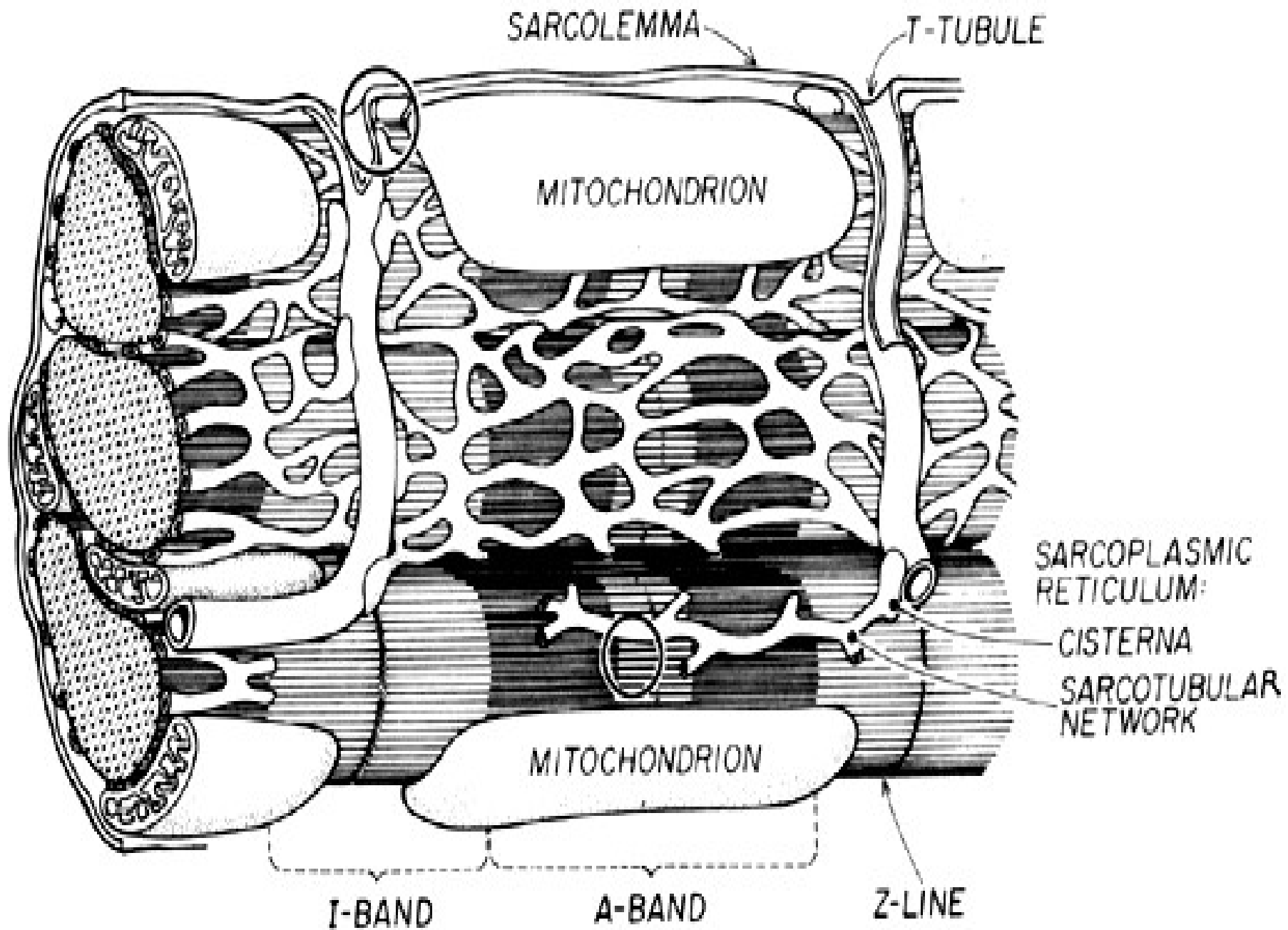
nexus

fascia adherens

Myofibril of cardiomyocyte

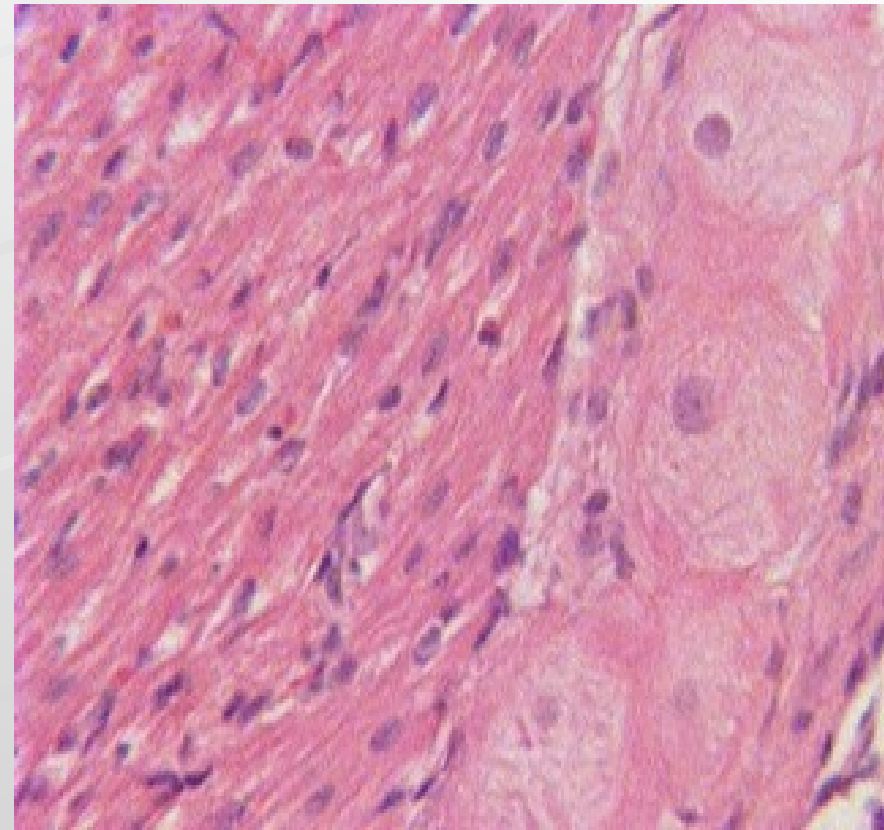
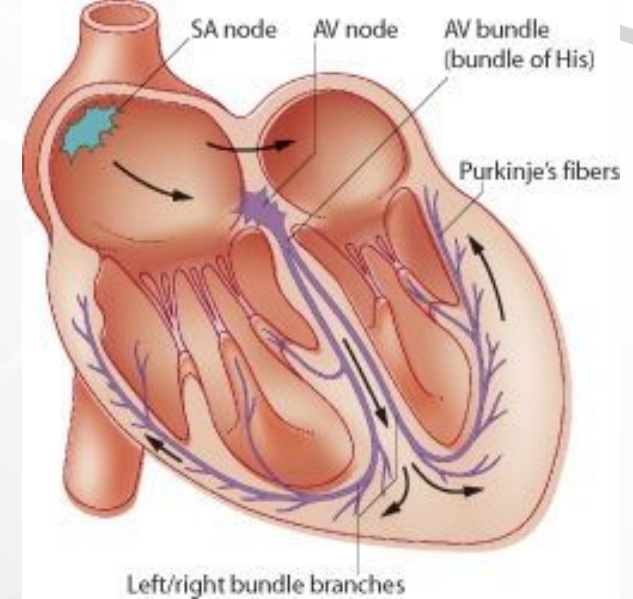
- ✦ Actin + myosin myofilaments
- ✦ Sarcomere
- ✦ Z-line
- ✦ M-line and H-zone
- ✦ I-band, A-band
- ✦ T-tubule + 1 cisterna = diad (around Z-line)





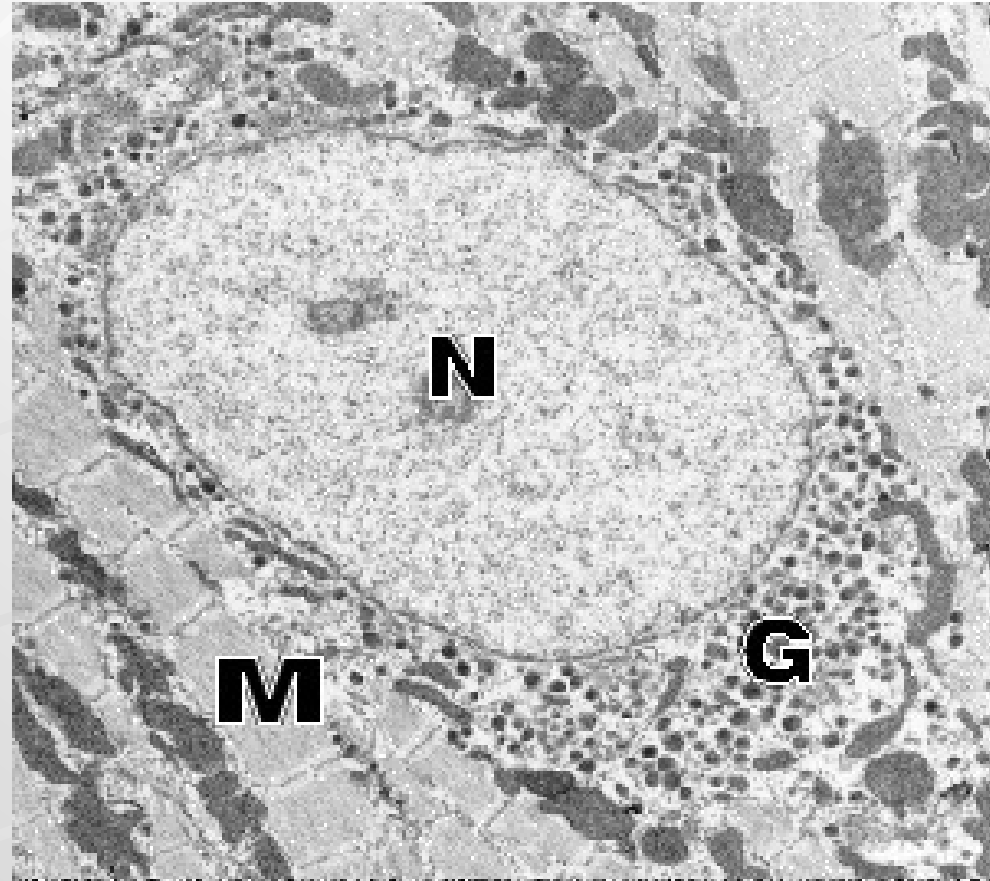
Purkinje's fibers

- ✚ are located in the inner layer of heart ventricle wall
- ✚ are specialized cells fibers that conduct an electrical stimuli or impulses that enables the heart to contract in a coordinated fashion
- ✚ numerous sodium ion channels and mitochondria, fewer myofibrils



Natriuretic hormone

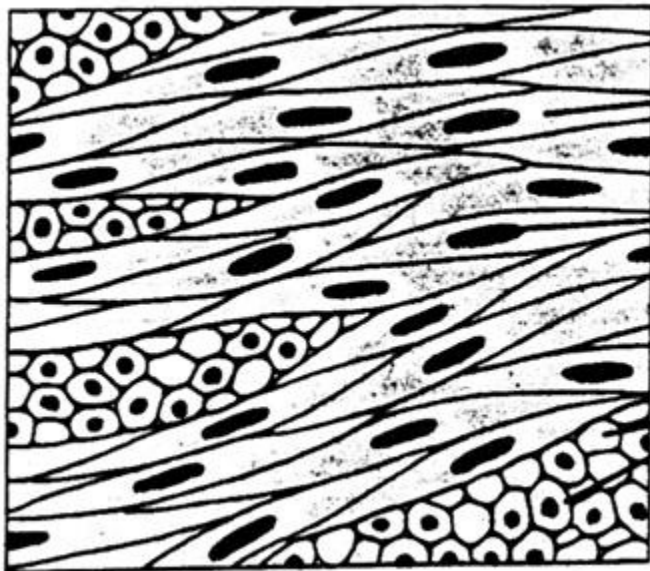
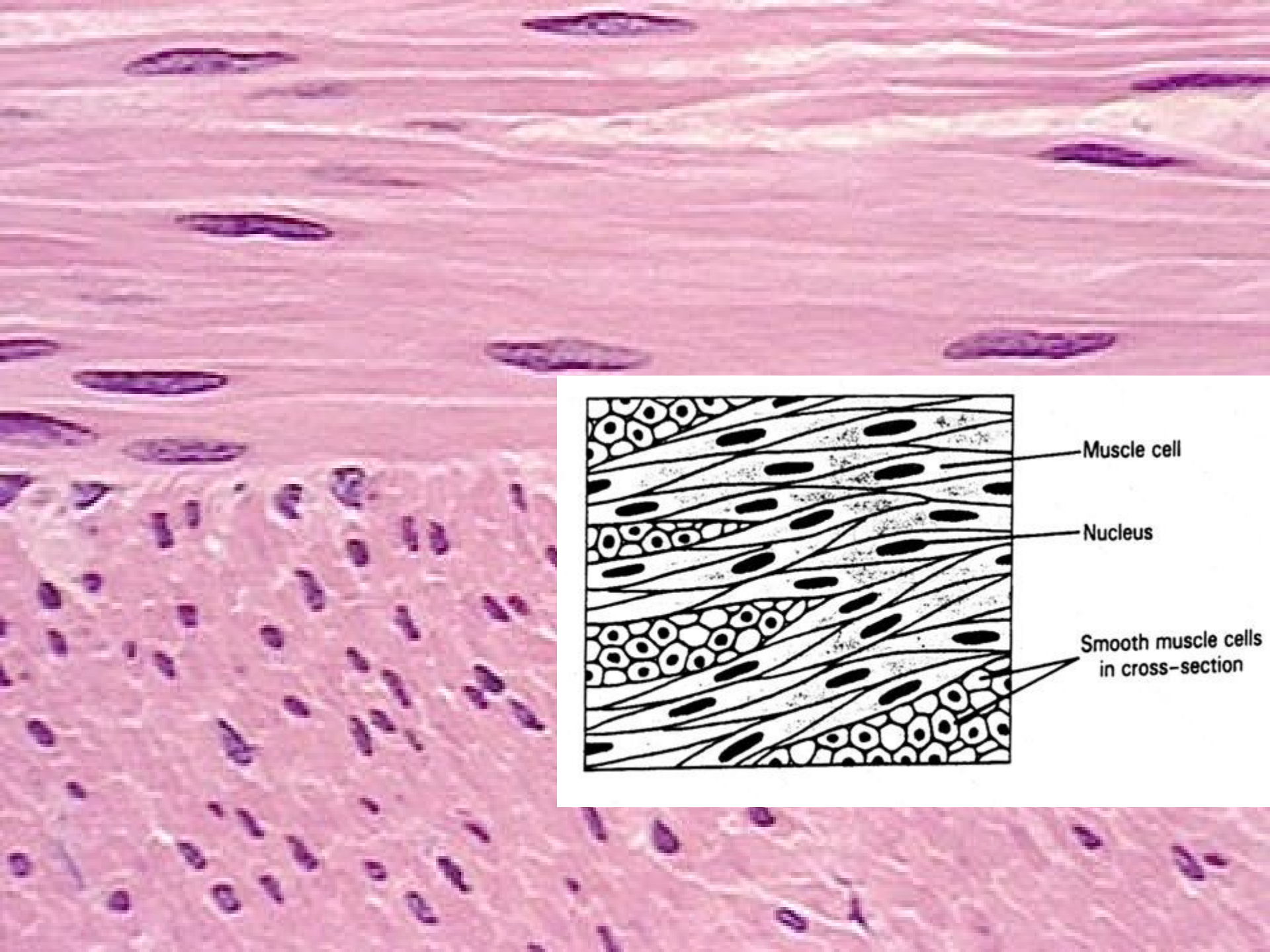
- ✦ in atrial granules in sarcoplasm of cardiomyocytes in ventricles
- ✦ 300 – 400 nm \emptyset
- ✦ this hormone is involved in regulation of volume of circulating blood



Electron microscopic view of atrial cardiocyte showing nucleus (N), myofibrils (M) and the storage site for ANF and BNP: atrial specific granules (G)

Smooth muscle tissue

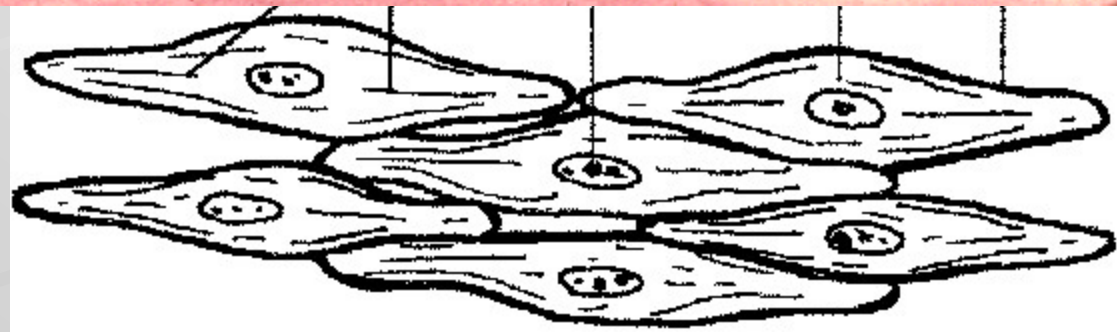
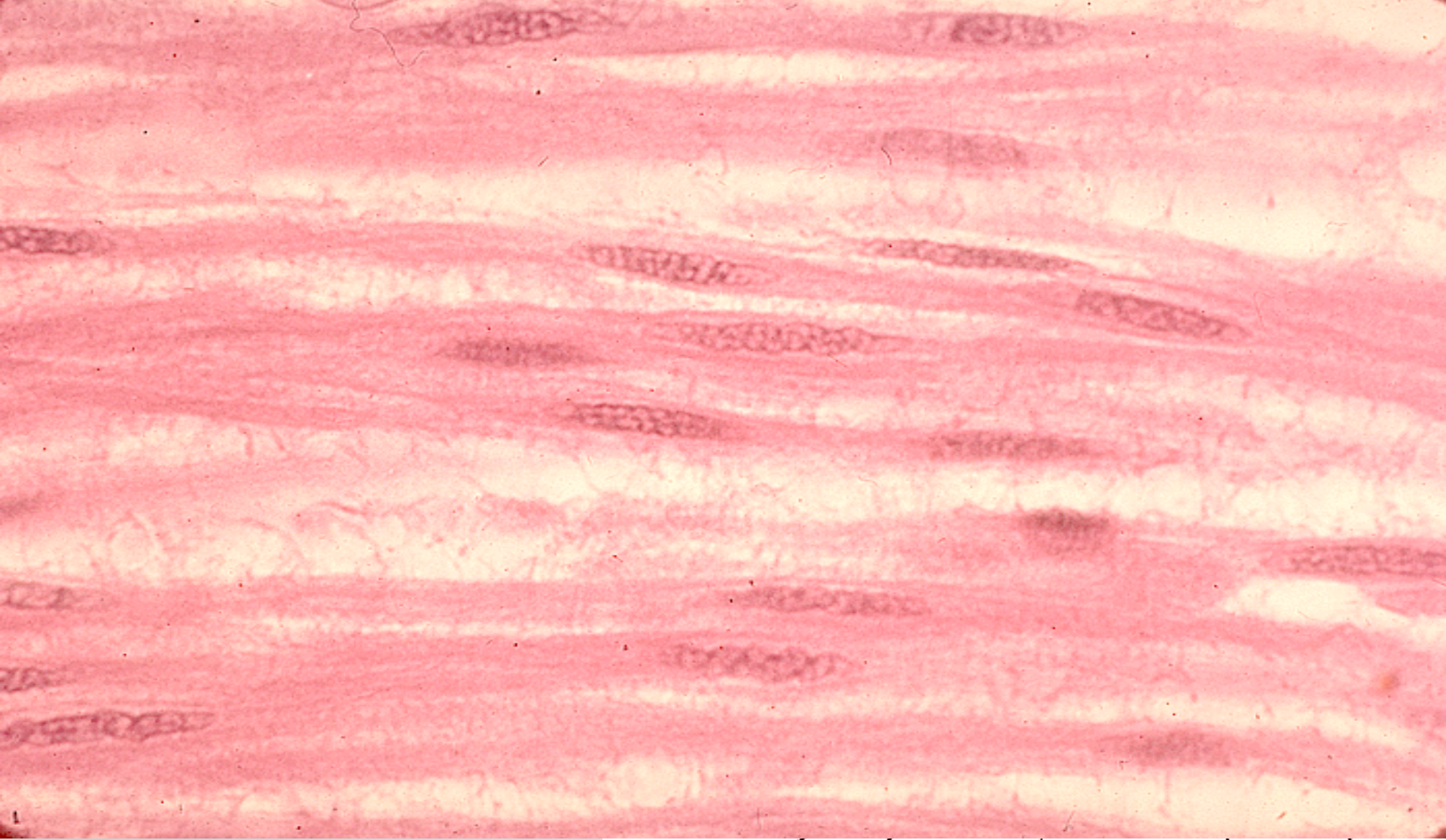
- ✚ spindle shaped cells (leiomyocytes) with myofilaments not arranged into myofibrils (no striation), 1 nucleus in the centre of the cell
- ✚ myofilaments form bands throughout the cell
- ✚ actin filaments attach to the sarcolemma by focal adhesions or to the **dense bodies** substituting Z-lines in sarcoplasm
- ✚ **calmodulin** (has function as troponin)
- ✚ sarcoplasmic reticulum forms only tubules
- ✚ **Caveolae** (have function as T-tubules)
- ✚ zonulae occludentes and nexuses connect cells

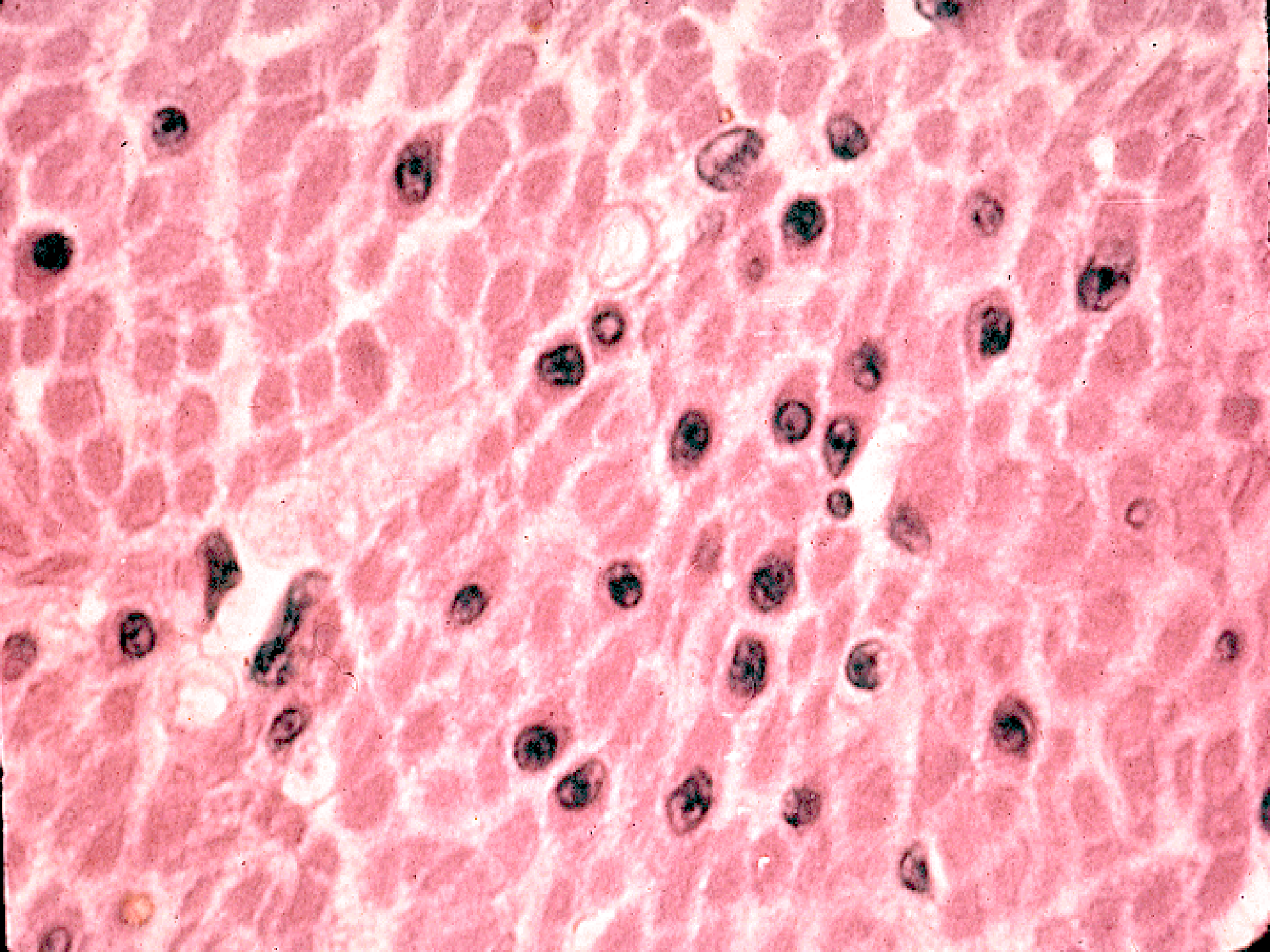


Muscle cell

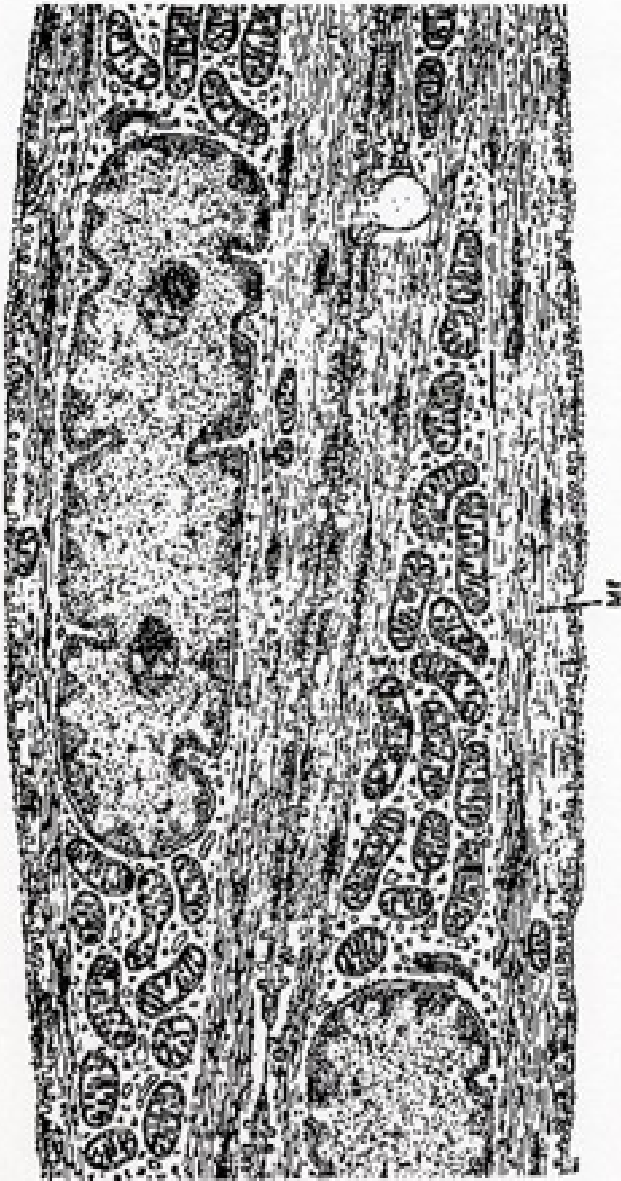
Nucleus

Smooth muscle cells
in cross-section





Leiomyocyte



Caveolae are equivalent to t-tubule and in their membrane ions channel are present to bring Ca needed for Contraction.

Caveolae are in contact with sarcoplasmic reticulum.

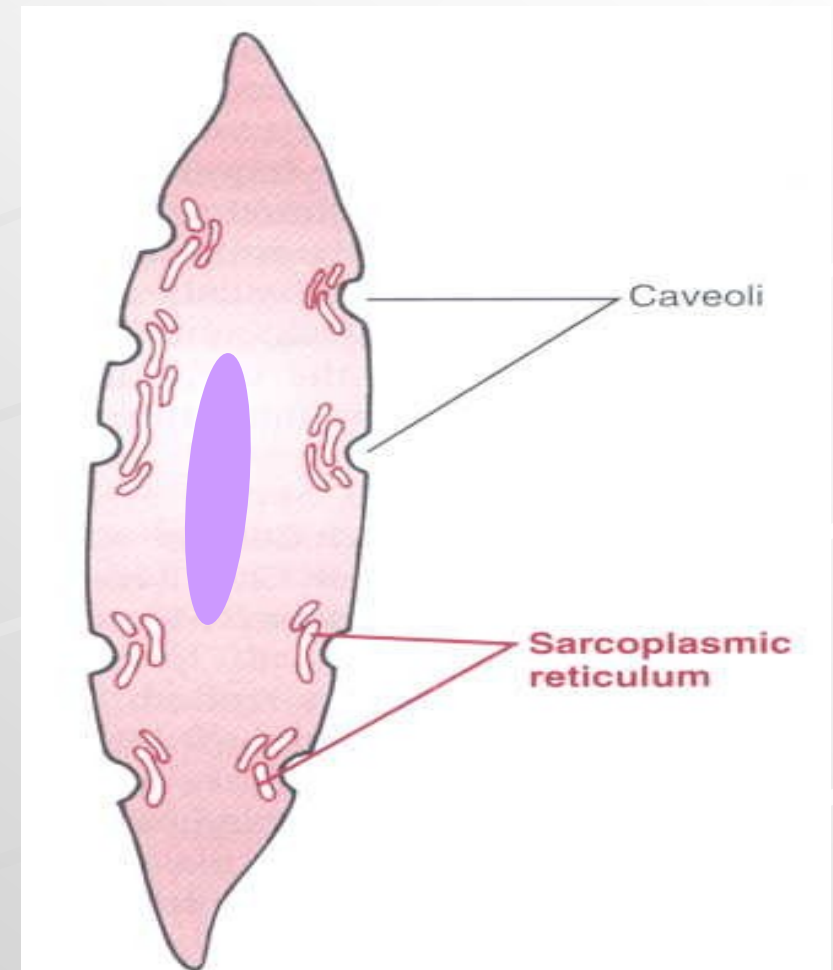
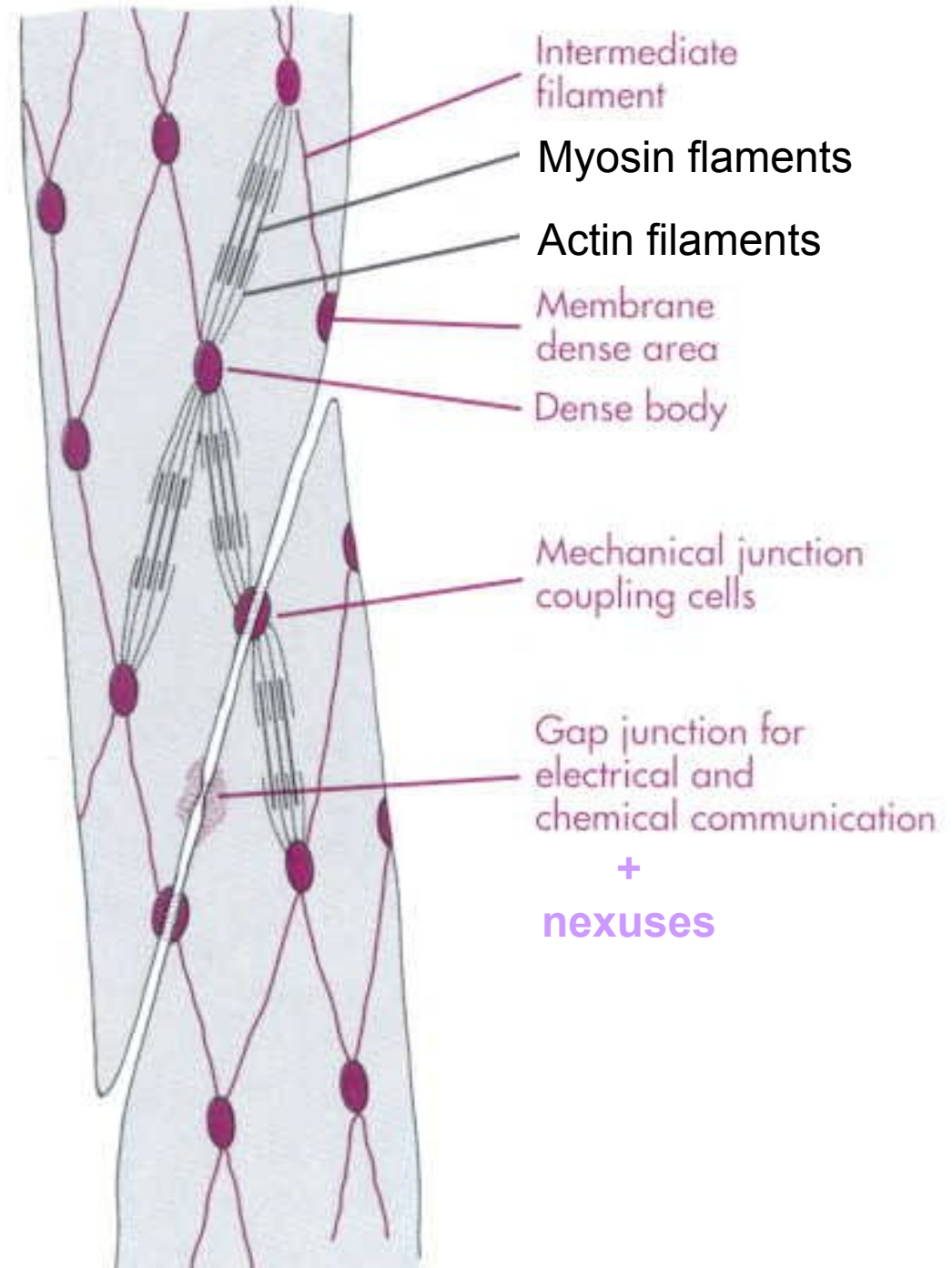
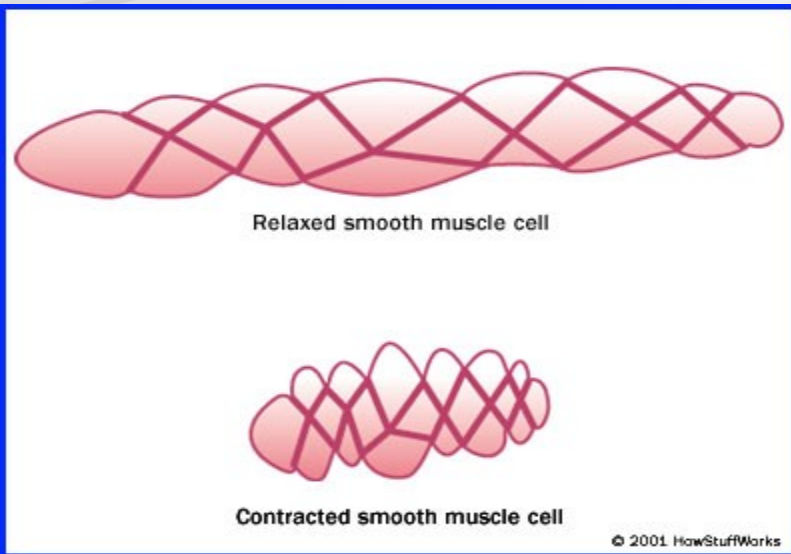


FIG. 10-2 E/M OF SMOOTH MUSCLE

Leiomyocyte: contractile filaments



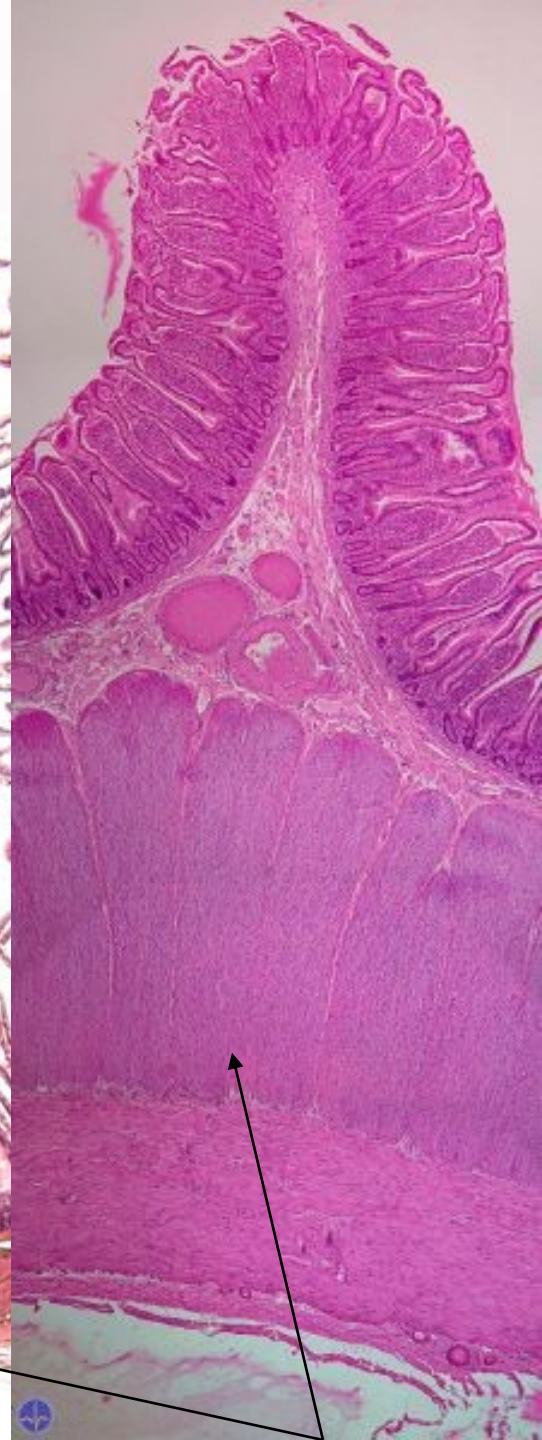
Muscle tissue

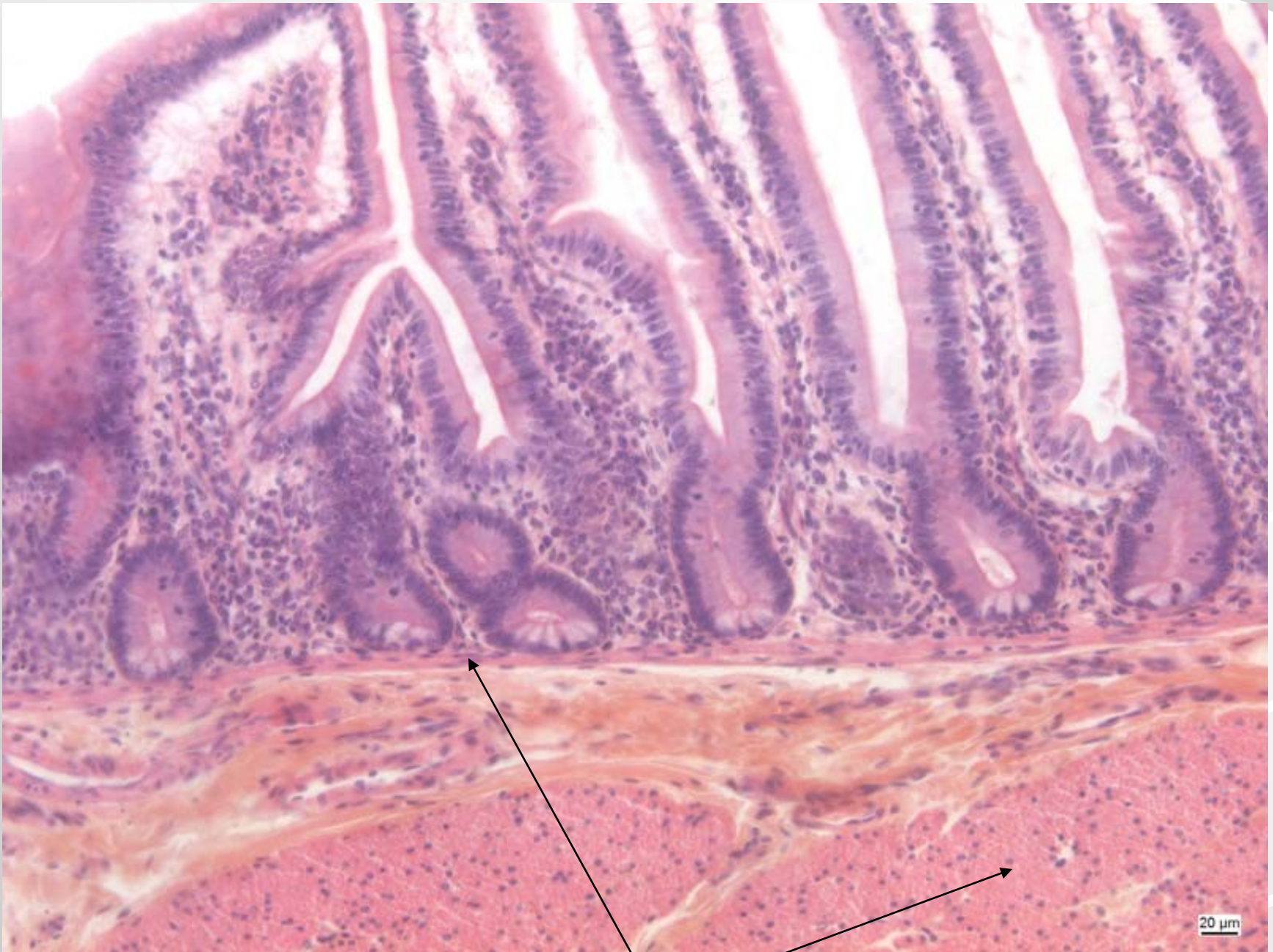
Slides:

- ✚ Skeletal muscle (2 – Apex linguae)
- ✚ Heart muscle (64, 65 – Myocardium)
- ✚ Smooth muscle (16 – Intestinum tenue)



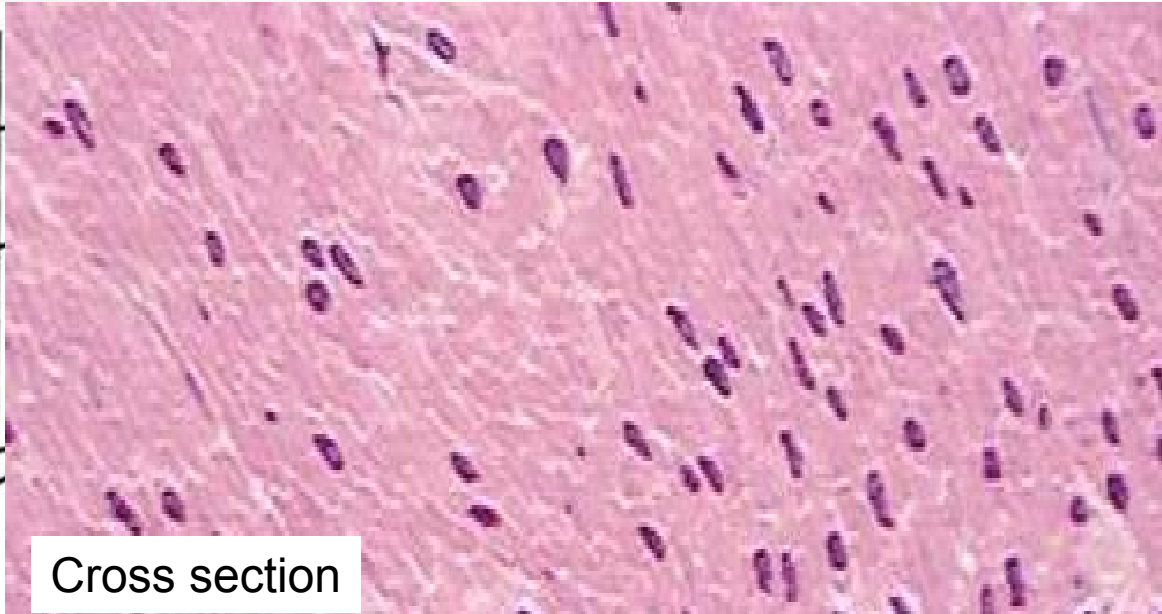
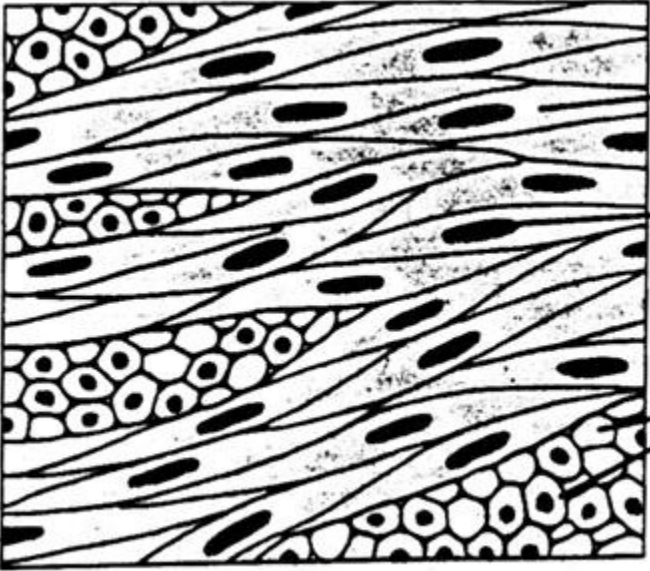
Smooth muscle



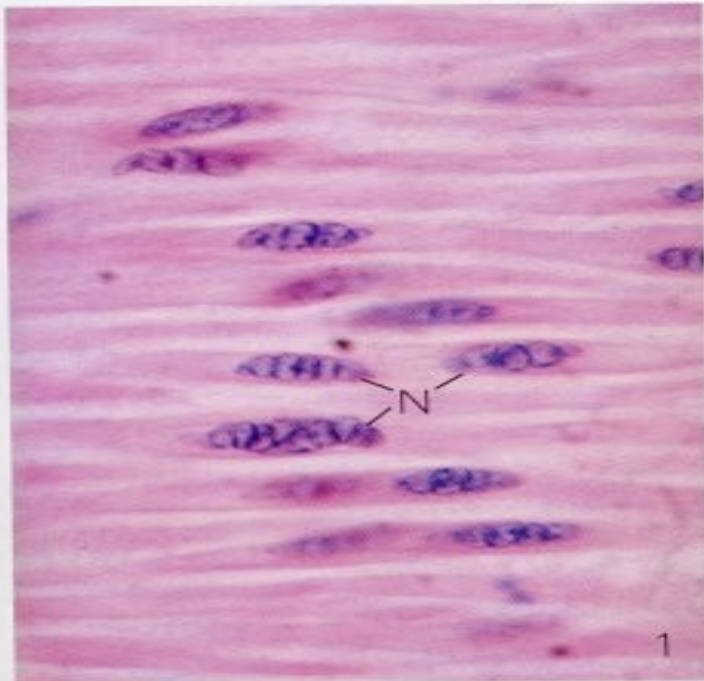


intestinum tenue – smooth muscle

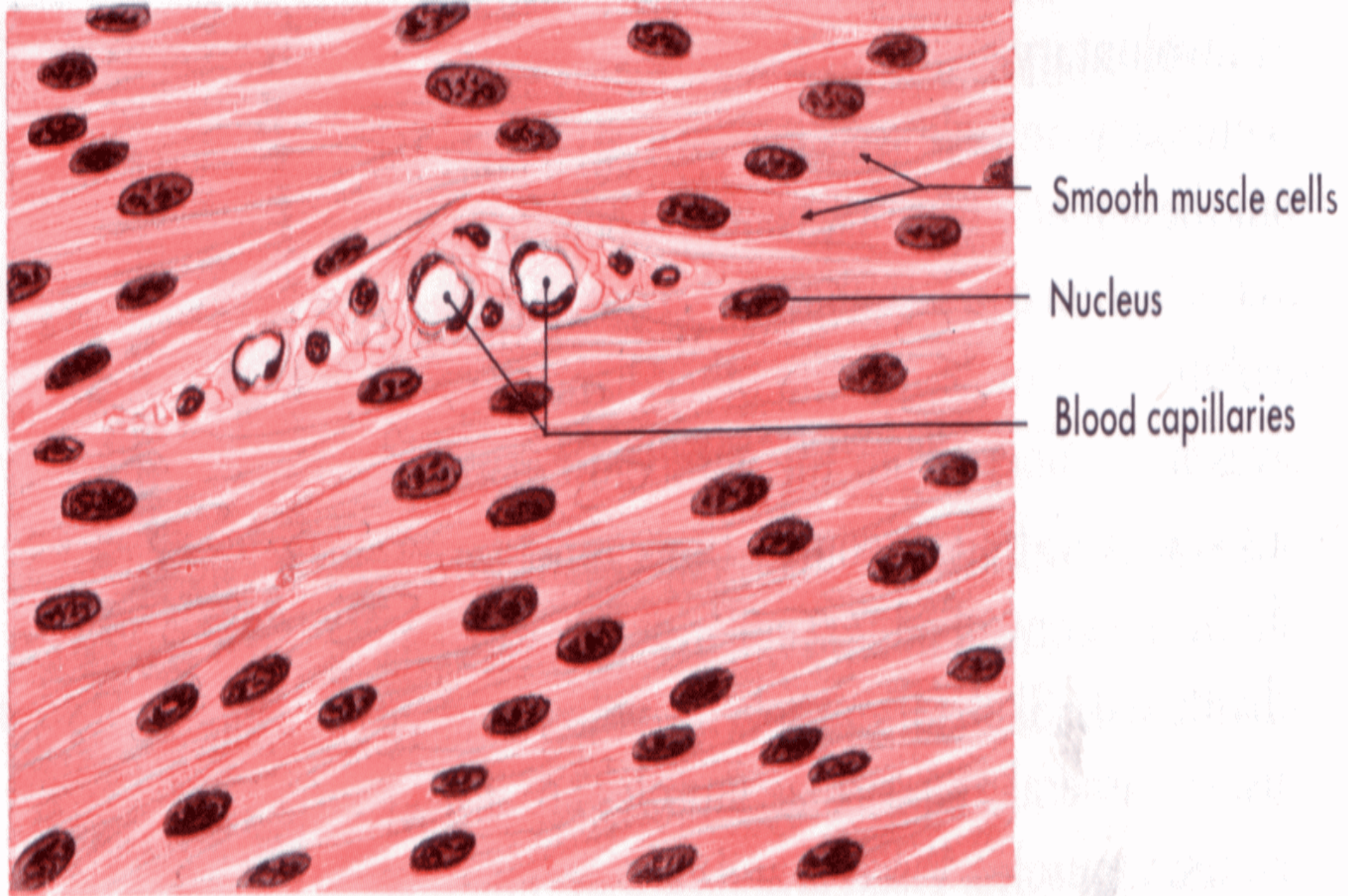
Leiomyocytes are arranged into layers of wall of hollow (usually tubular) organs



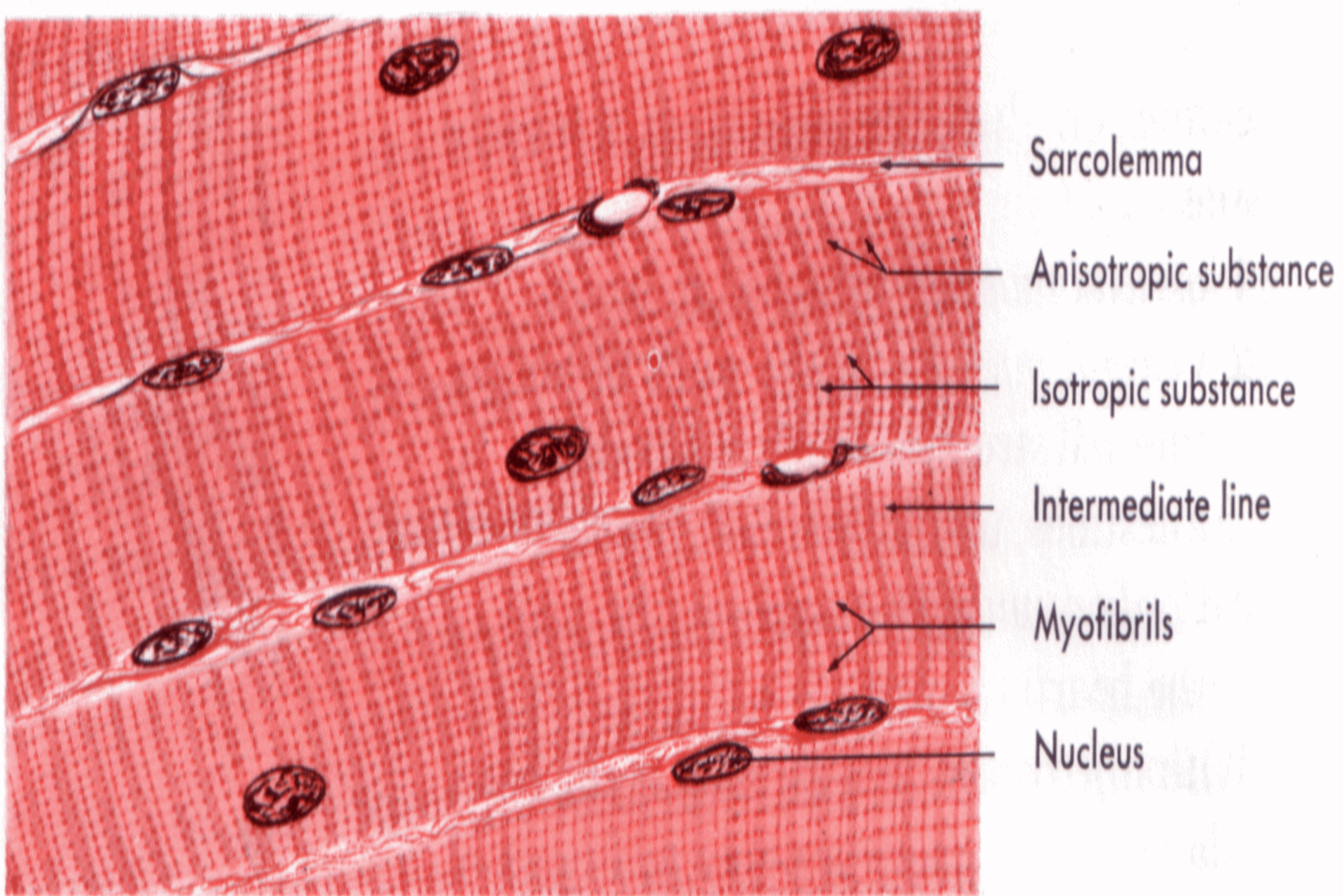
Cross section



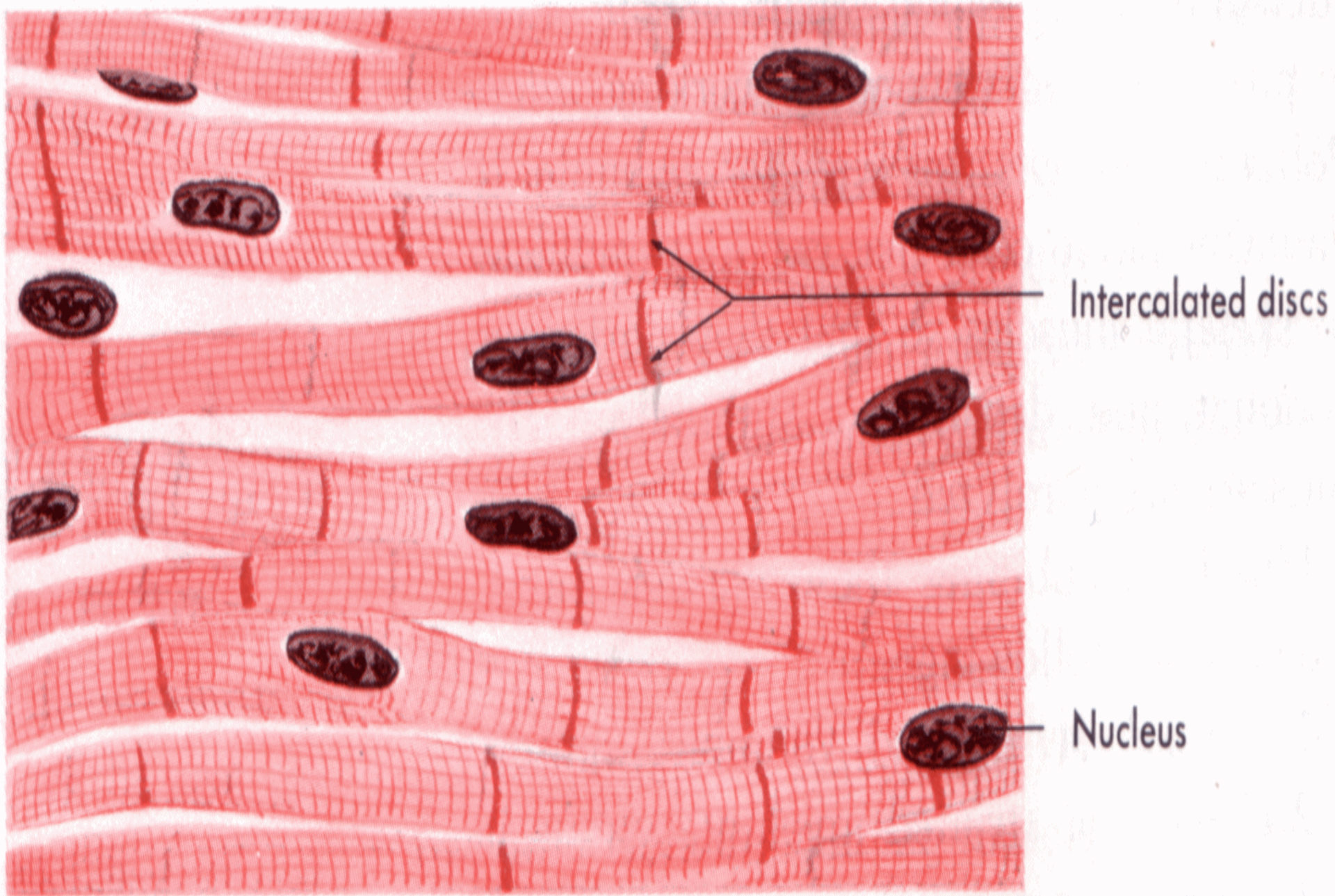
Longitudinal section



Visceral or nonstriated (smooth) involuntary muscle tissue.



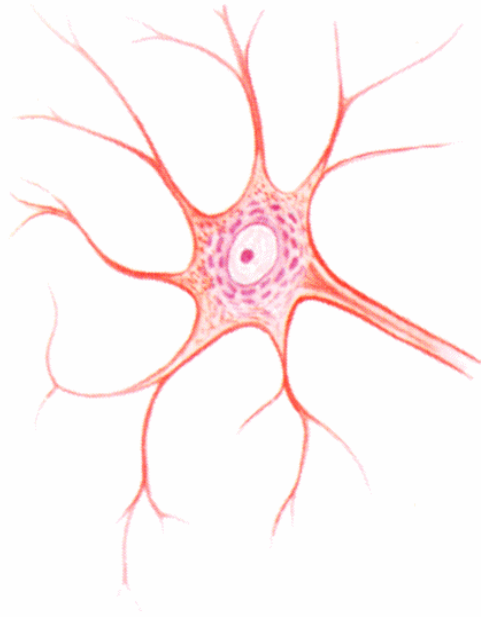
Skeletal or striated voluntary muscle tissue.



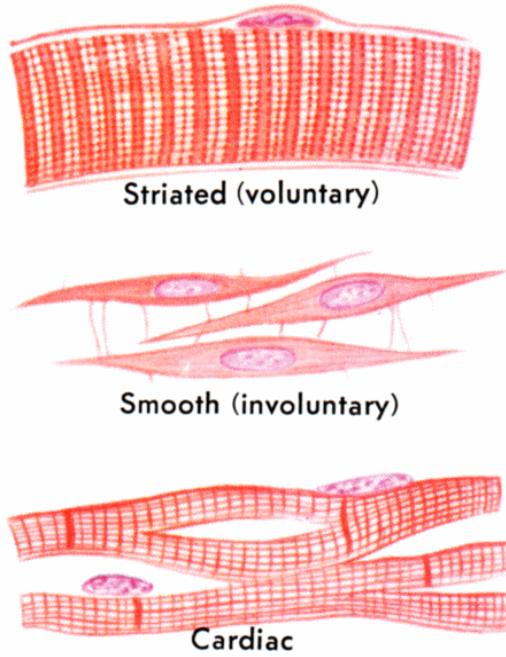
Cardiac or striated involuntary muscle tissue.

TYPES OF CELLS

NERVE CELL



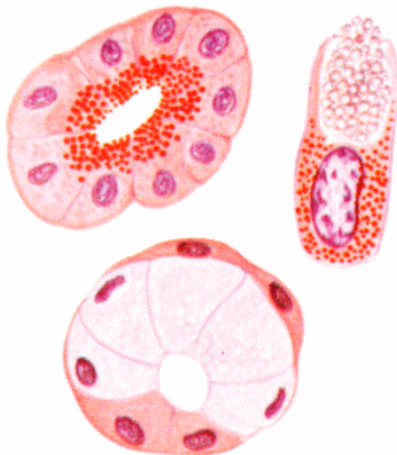
MUSCLE CELLS



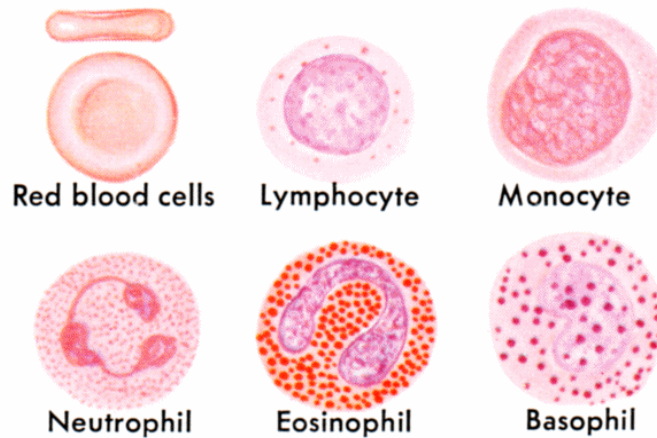
BONE CELL



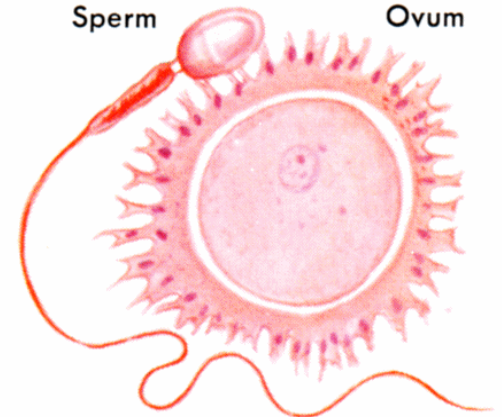
GLAND CELLS



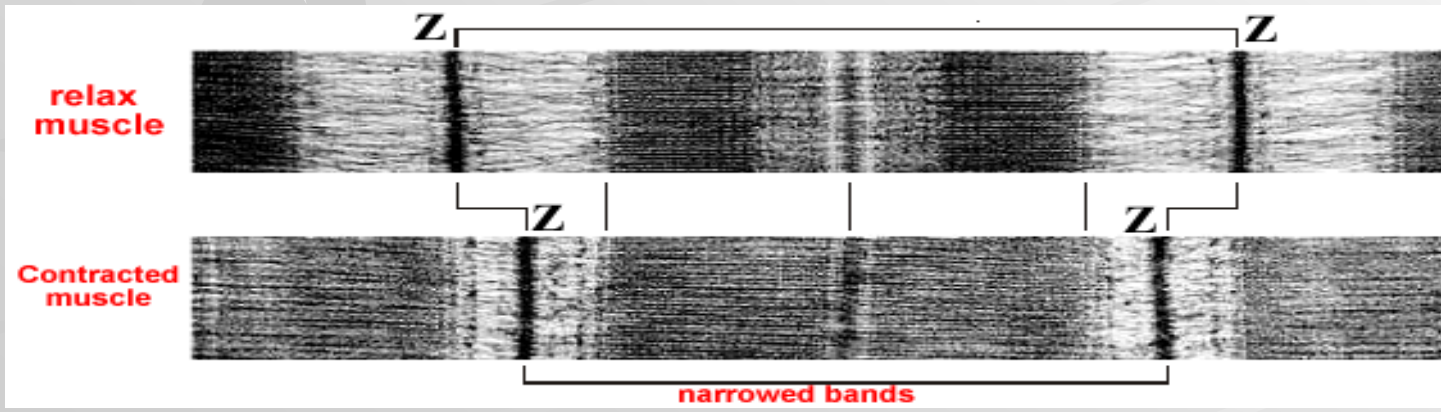
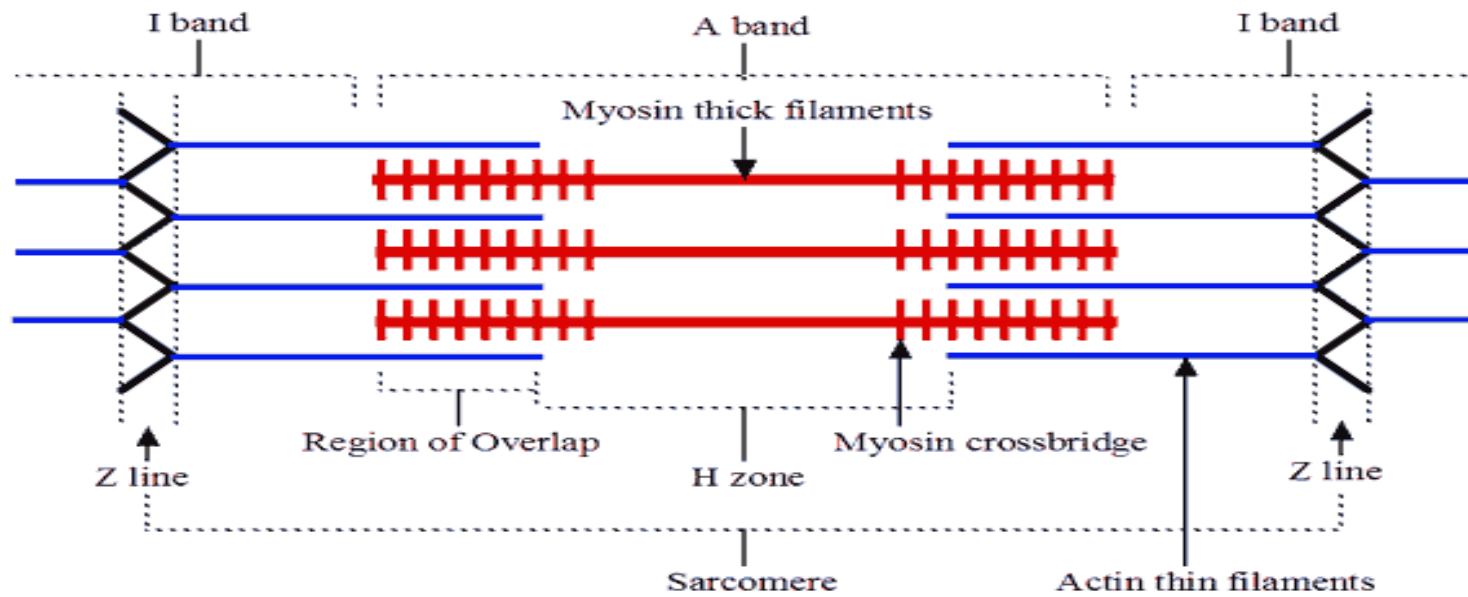
BLOOD CELLS

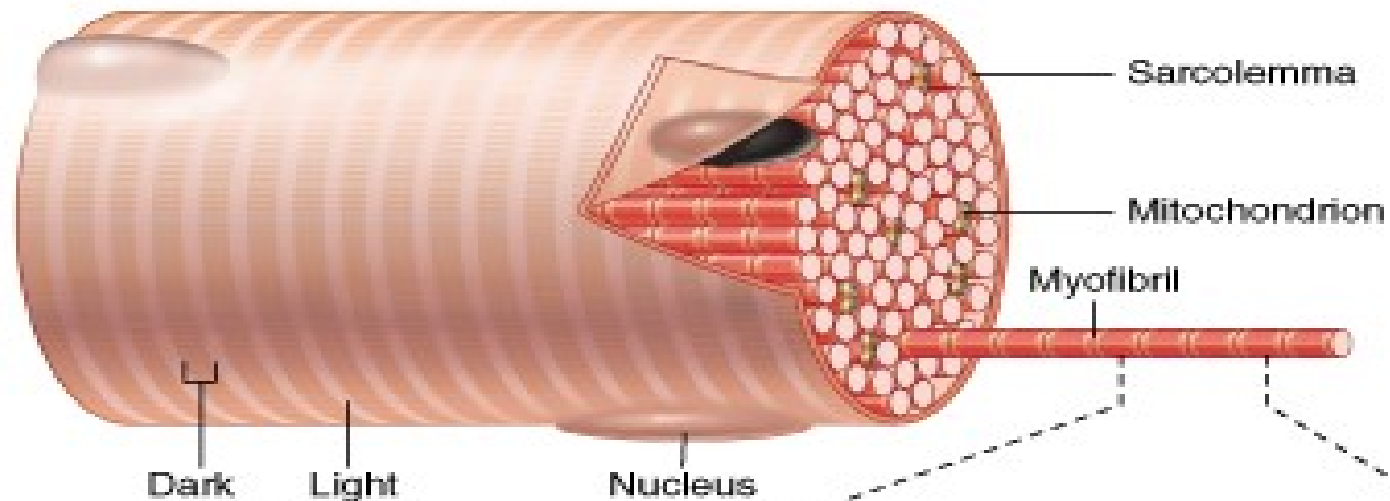


Sperm **Ovum**

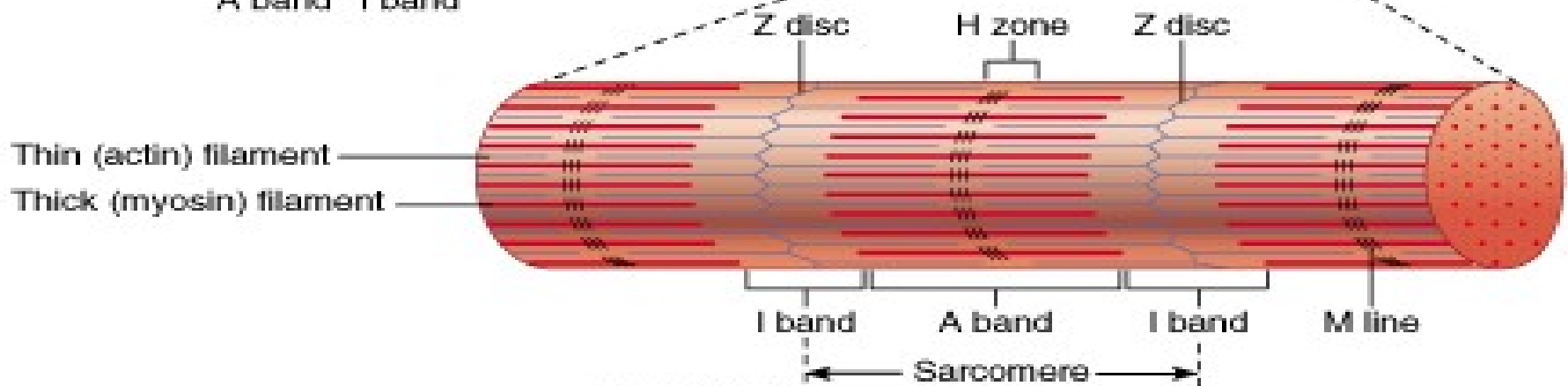


REPRODUCTIVE CELLS

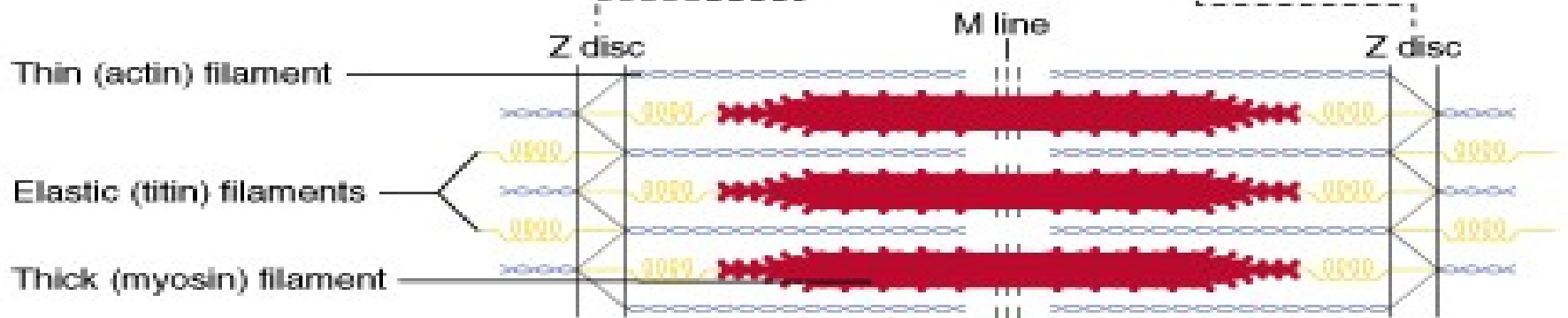




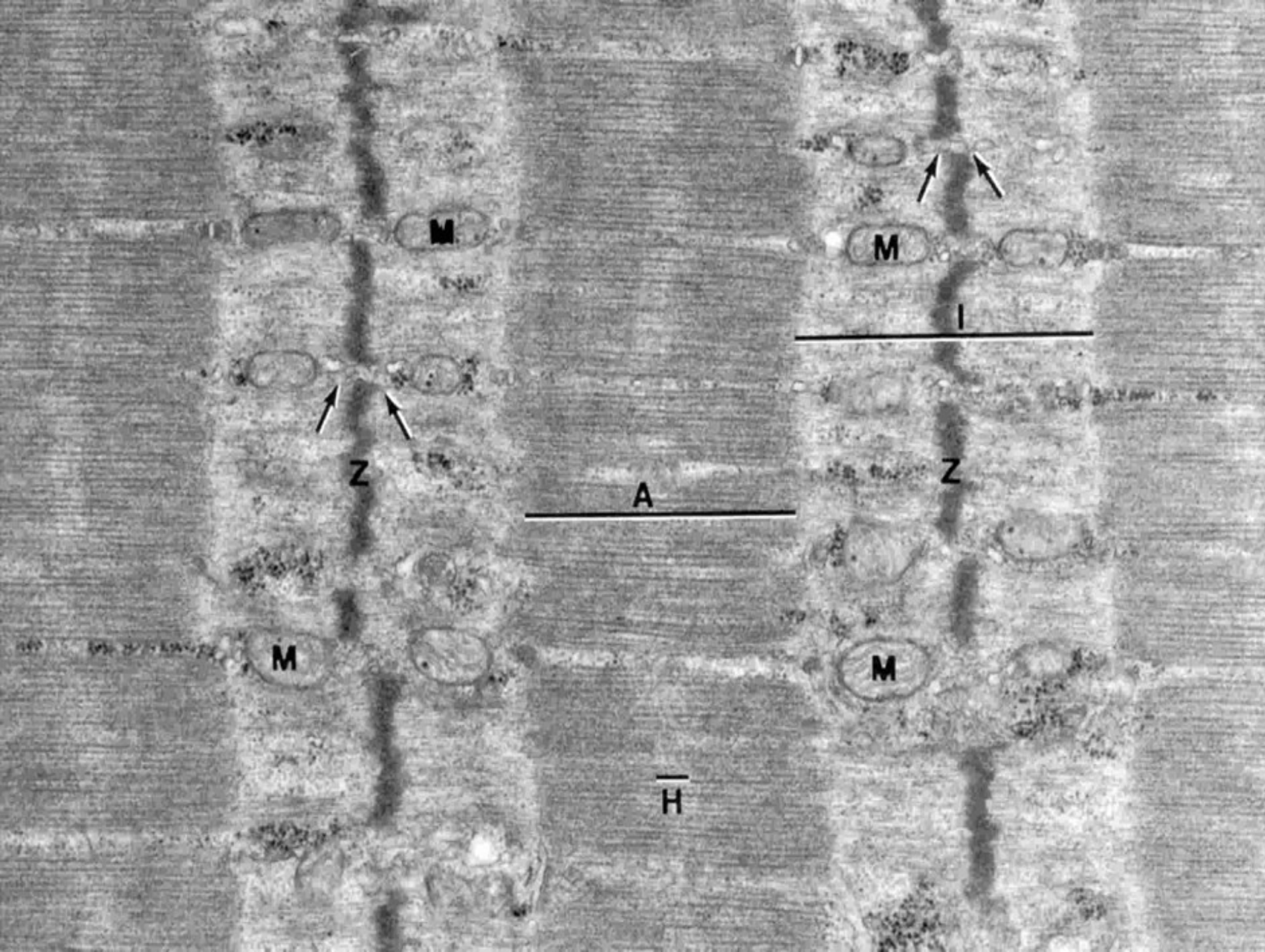
(b)

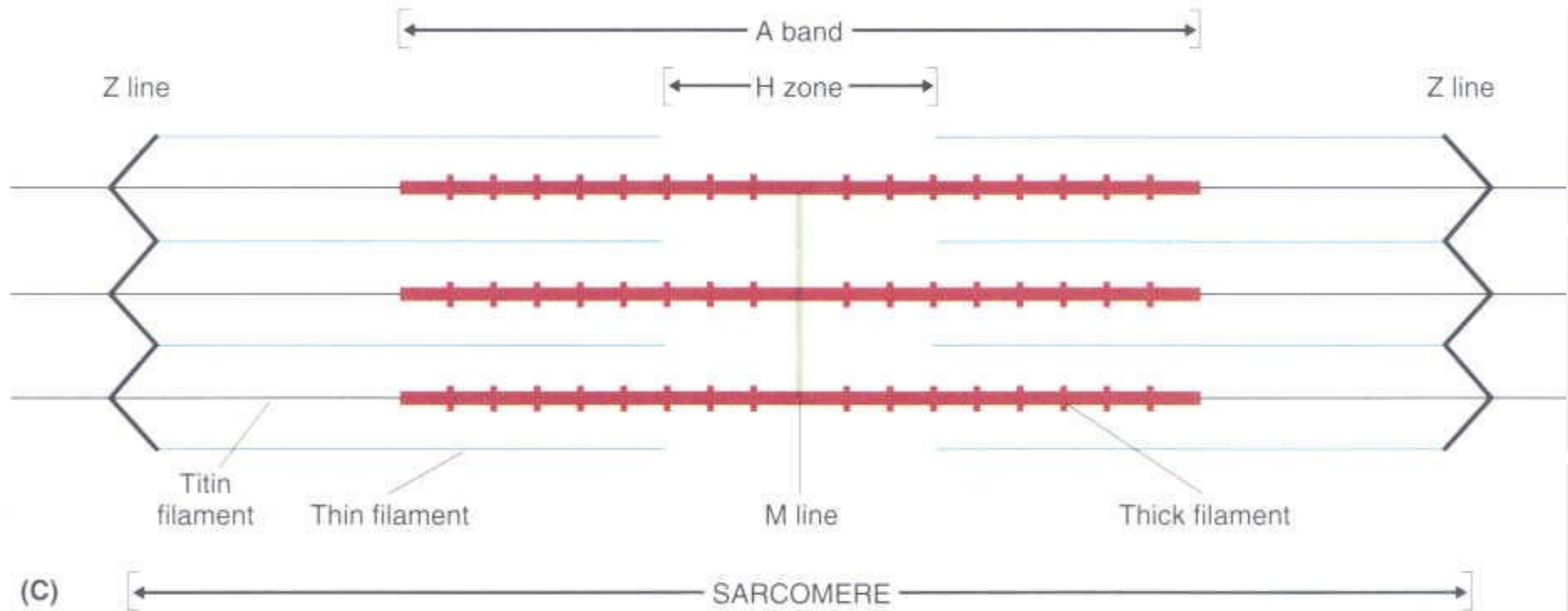
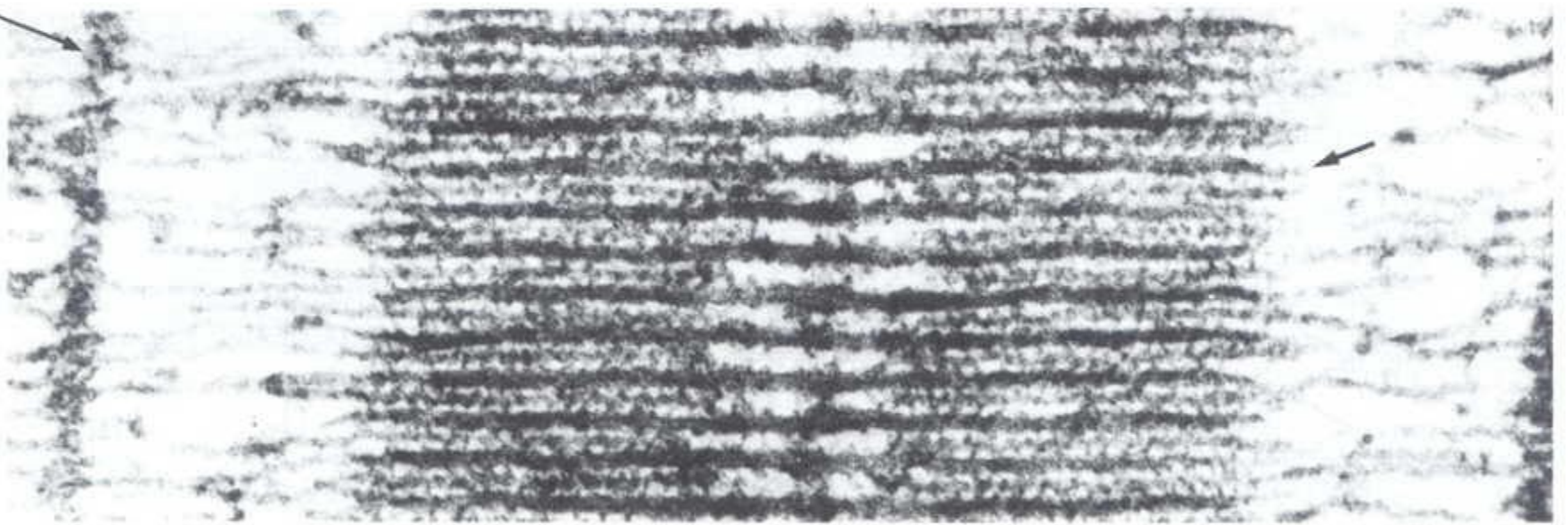


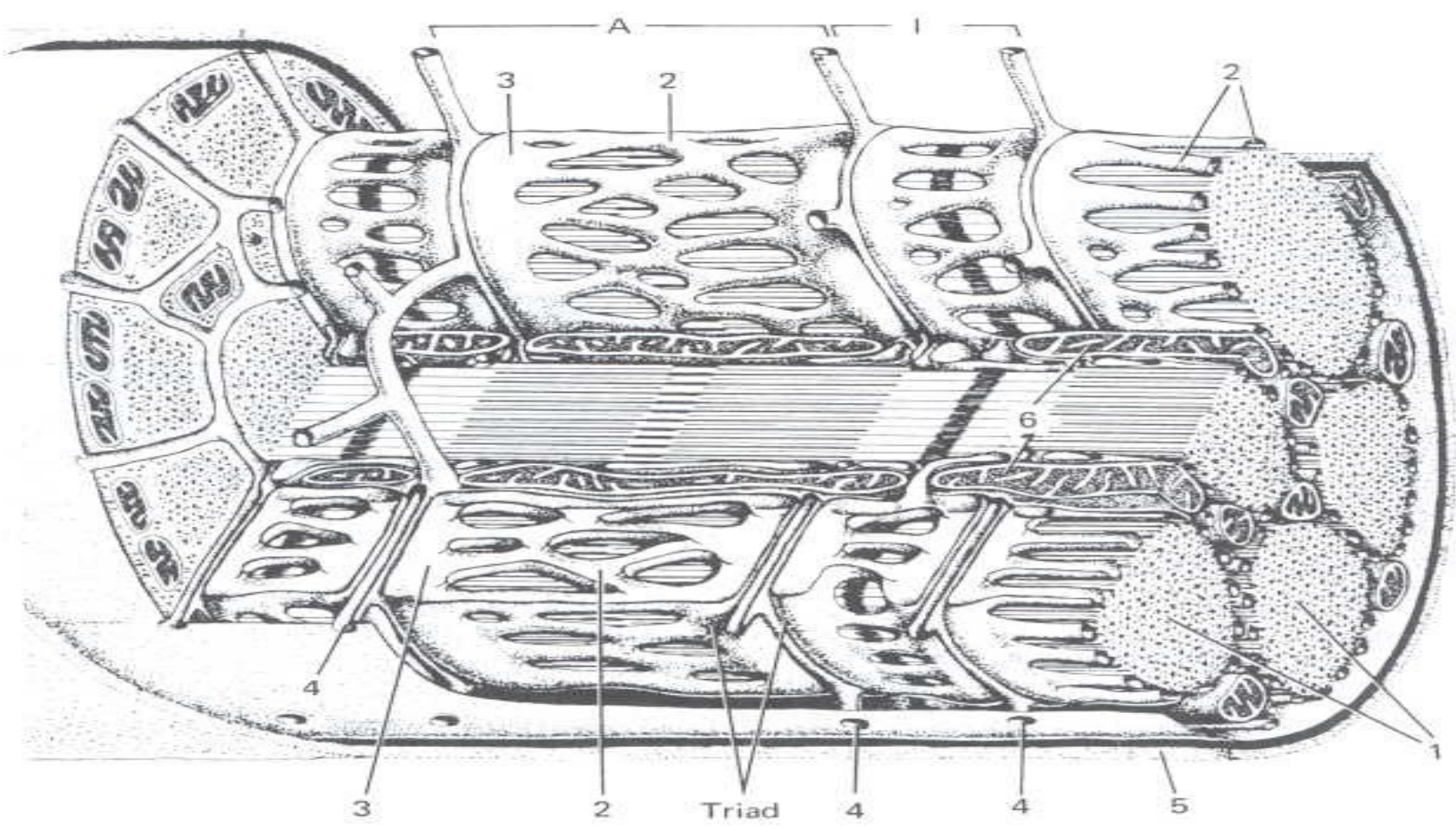
(c)

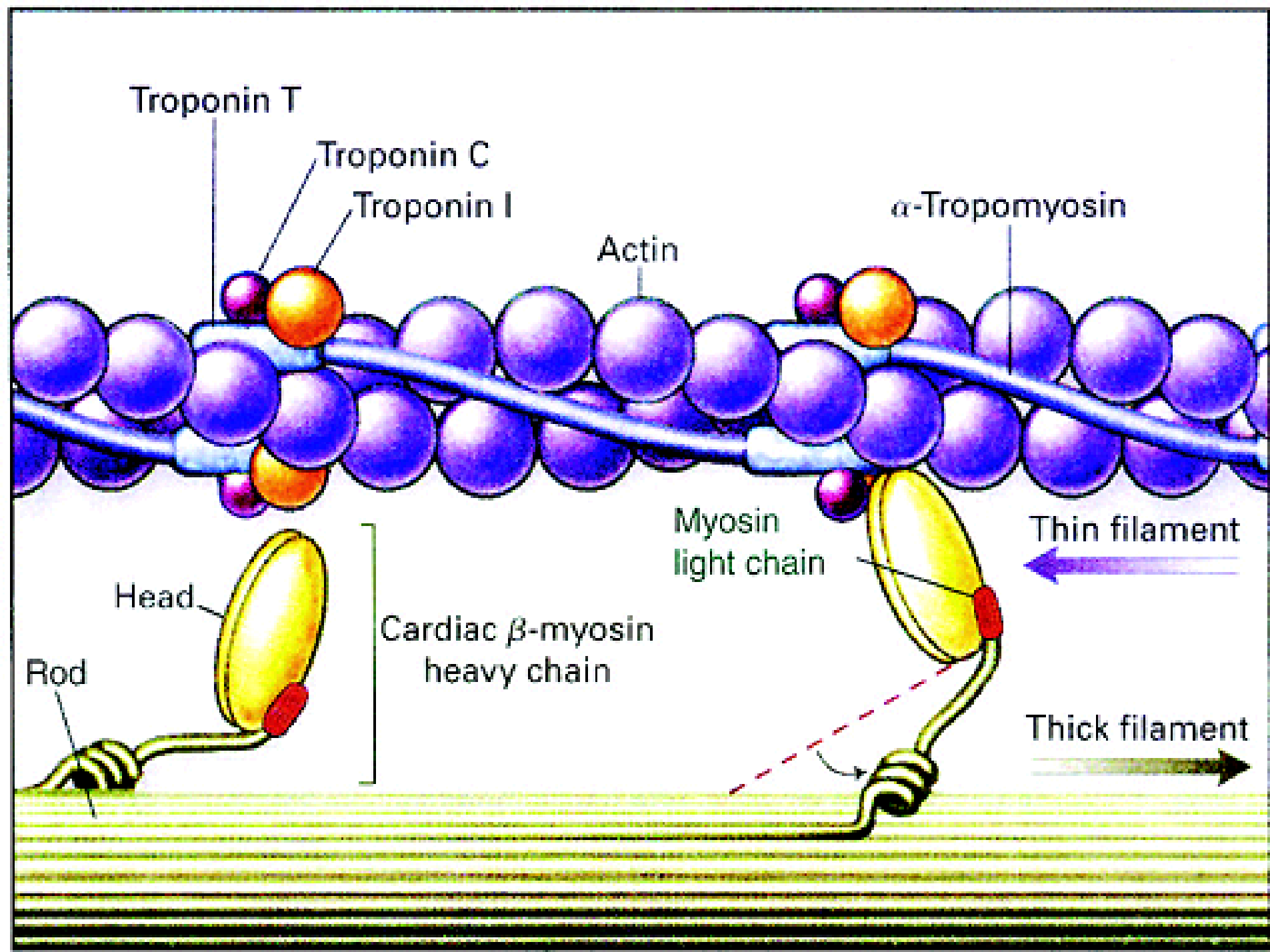


(d)

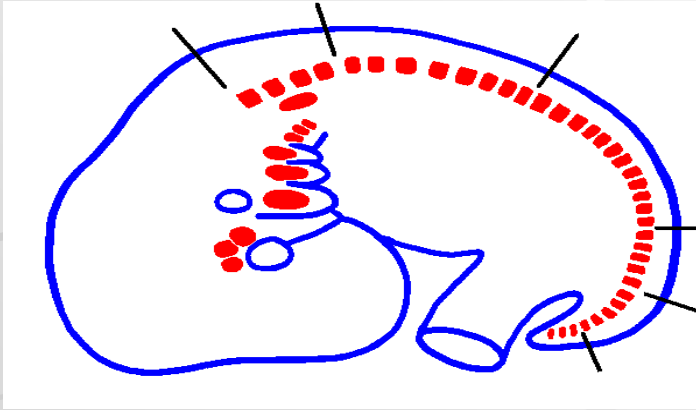






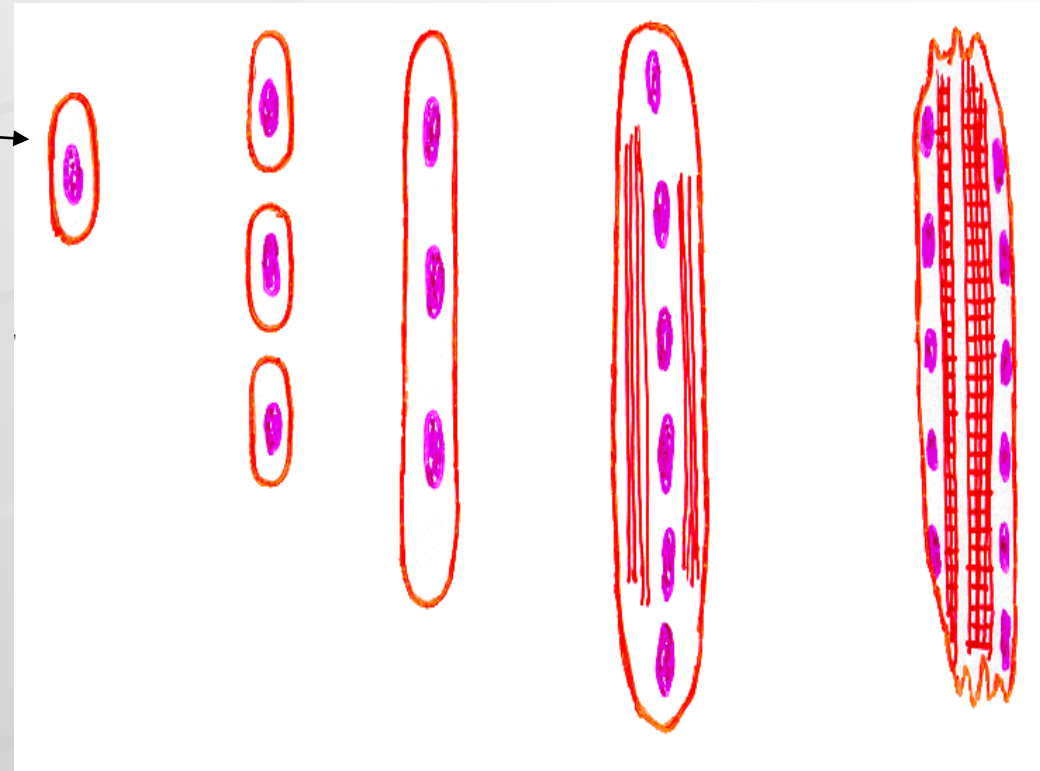


Development of rhabdomyocyte



Origin of skeletal muscle:
paraaxial mesoderm – somites – myotomes

Fusion of myoblasts
(primitive muscle cells
in myotomes) into
multinucleated
muscle fiber



Development of cardiomyocytes and leiomyocytes

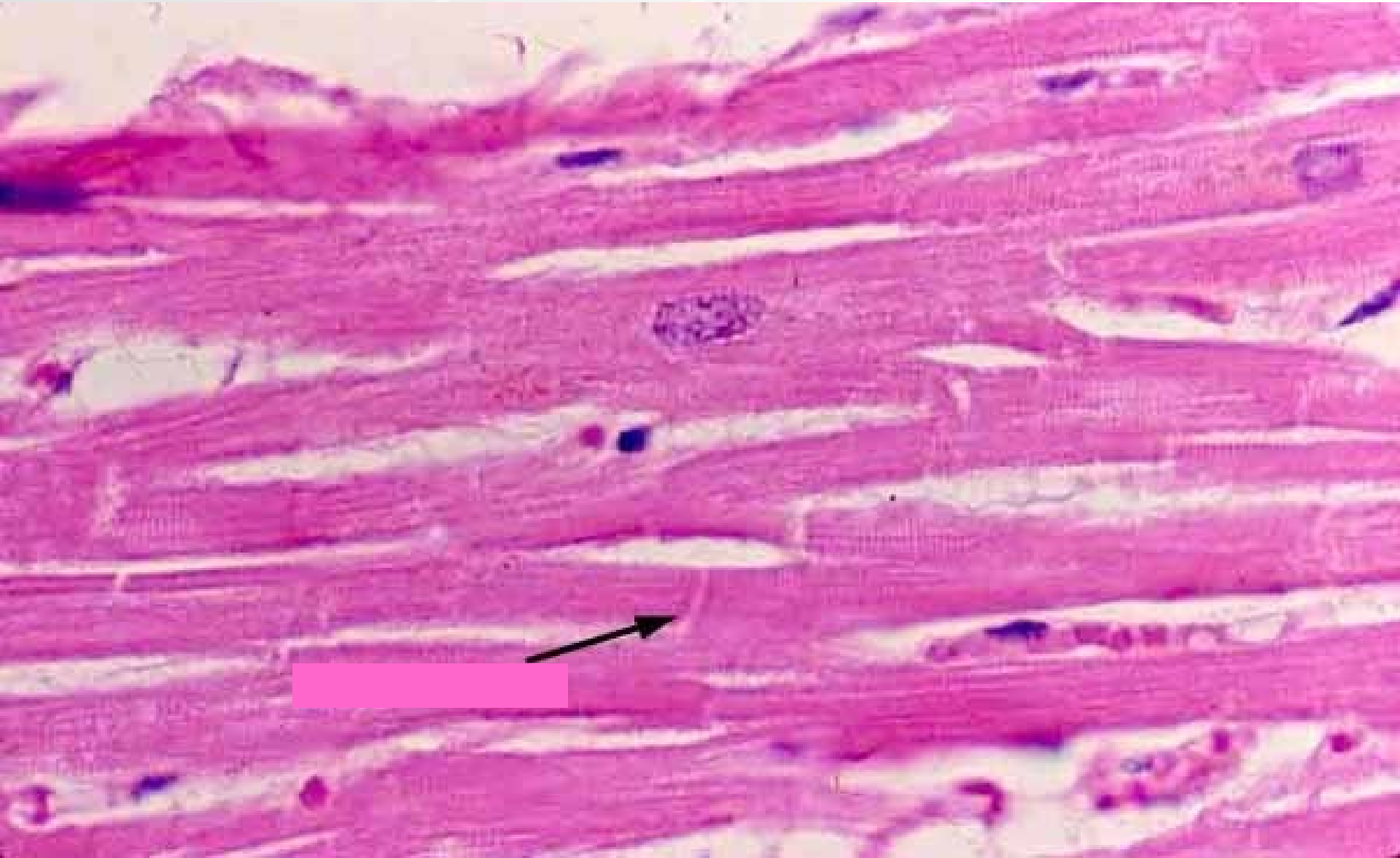


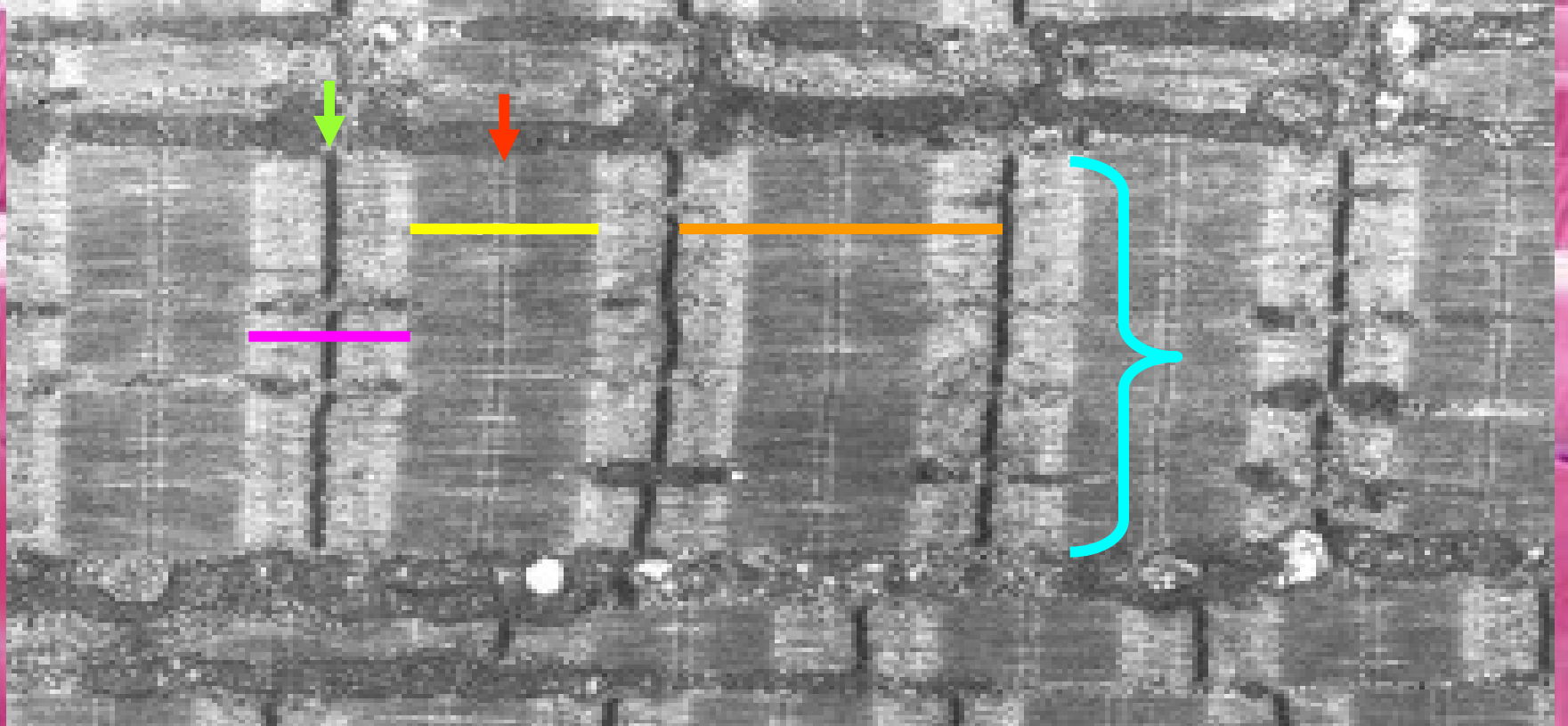
Practice and Quiz

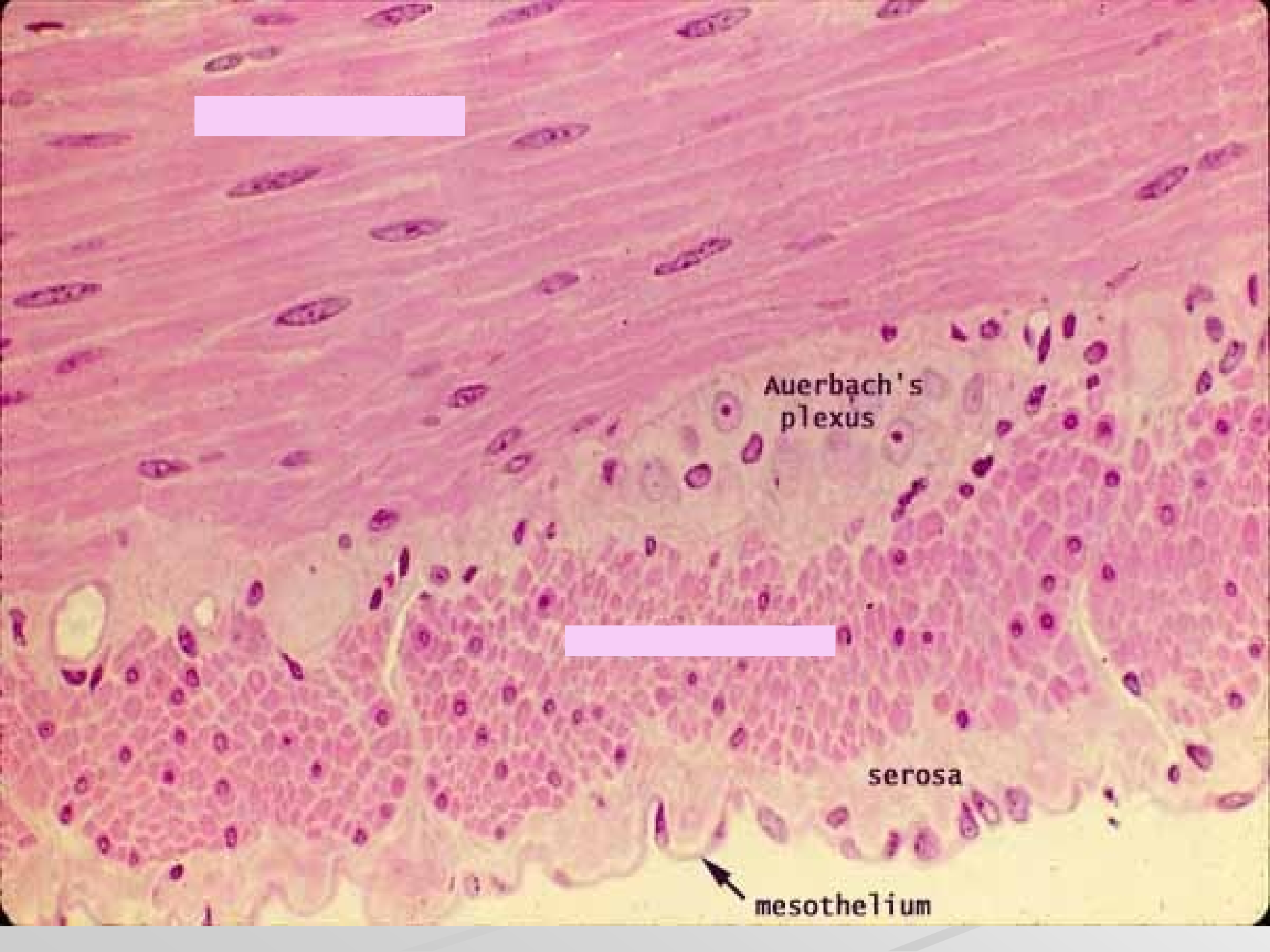


- ✦ Rhabdomyocyte
- ✦ Endomysium
- ✦ Sarcomere
- ✦ T-tubule
- ✦ Troponin
- ✦ Myofilament
- ✦ Myofibril
- ✦ Sarcoplasmic reticulum
- ✦ Actin
- ✦ Sarcolemme
- ✦ Perimysium

- Leiomycocyte
- Myosin
- Epimysium
- Triad
- Intercalated disc
- A-band
- Cardiomyocyte
- M-line
- Myosin
- Fascia adherens
- Motor-end-plate



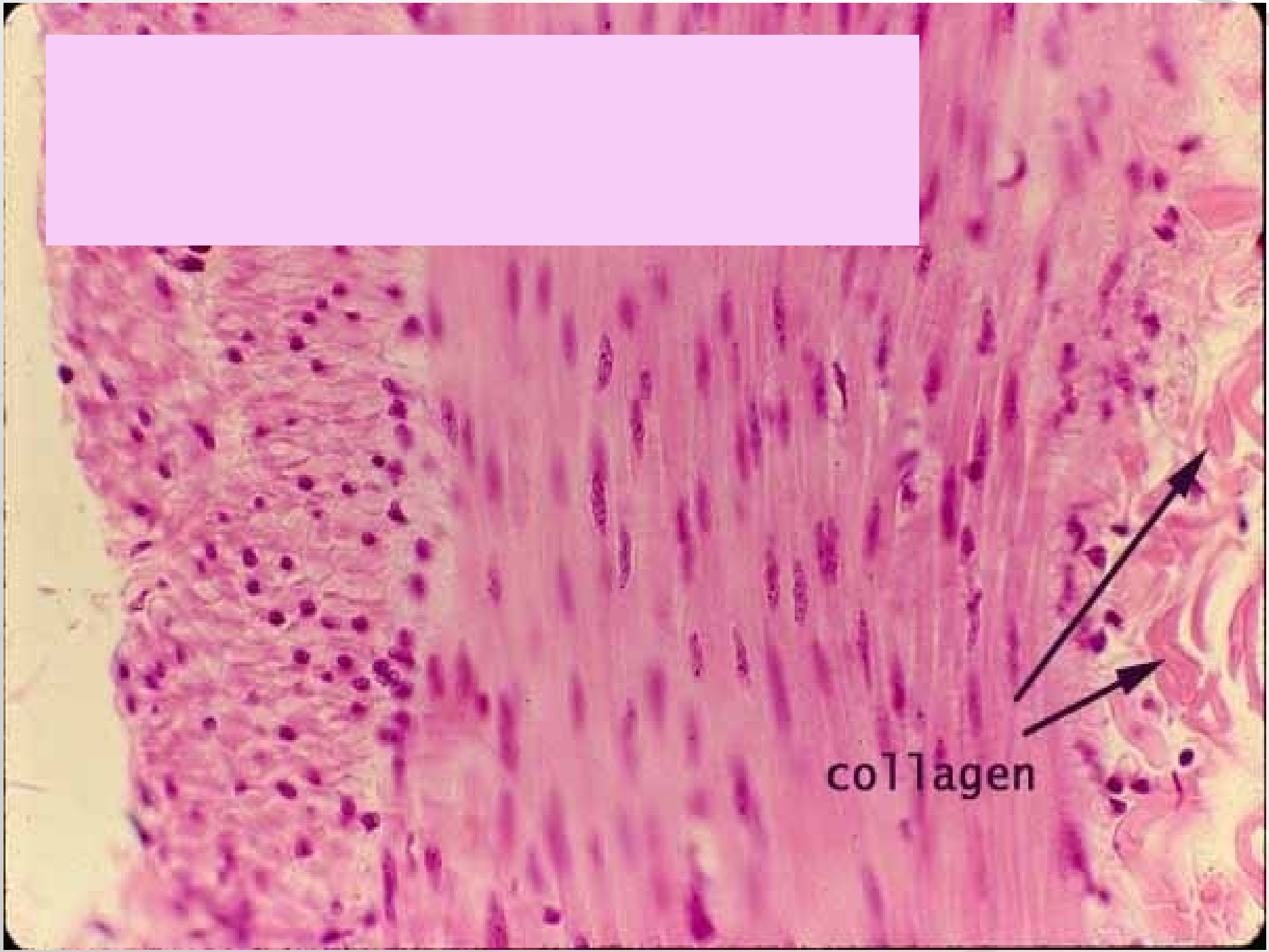




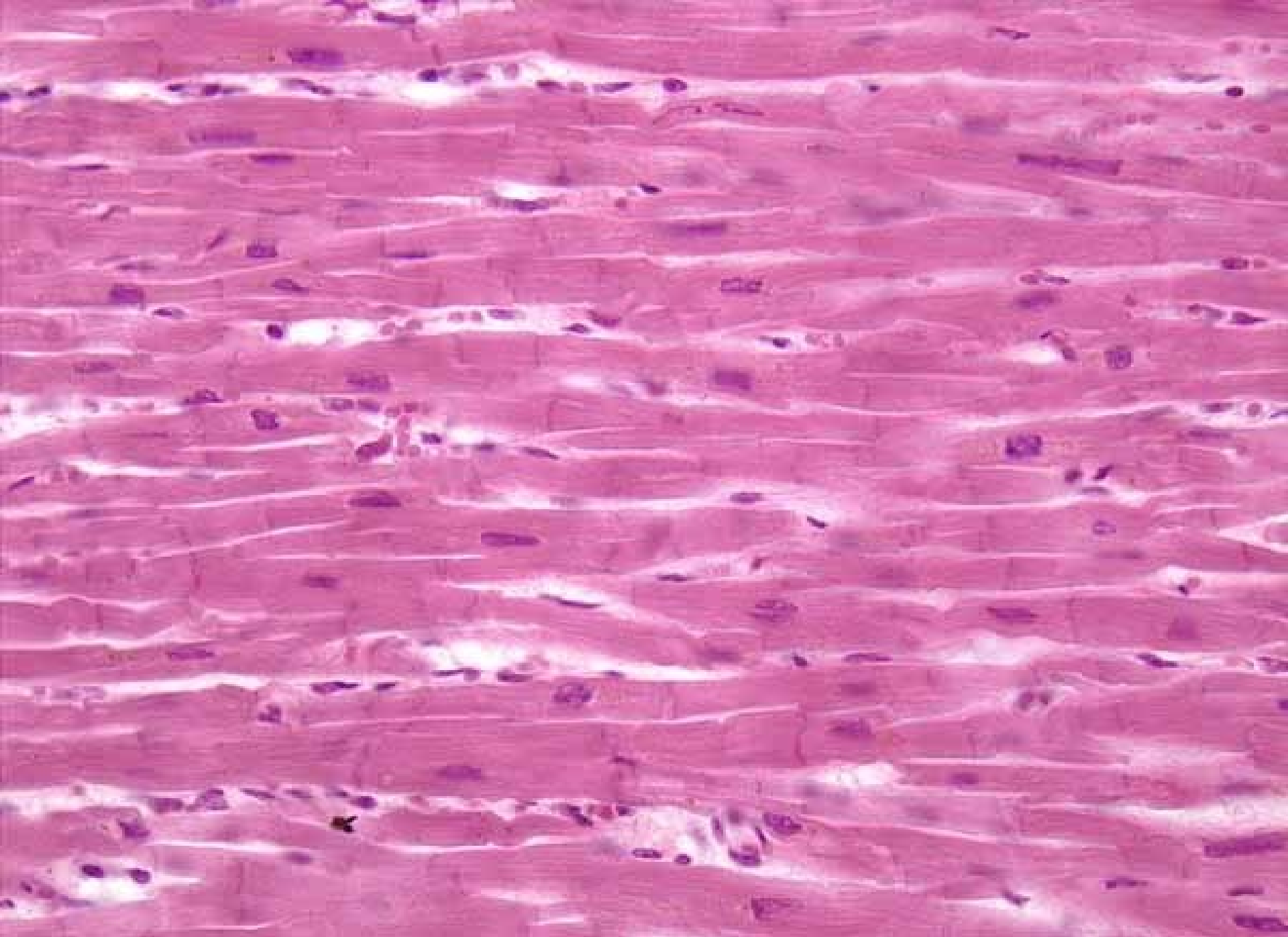
Auerbach's
plexus

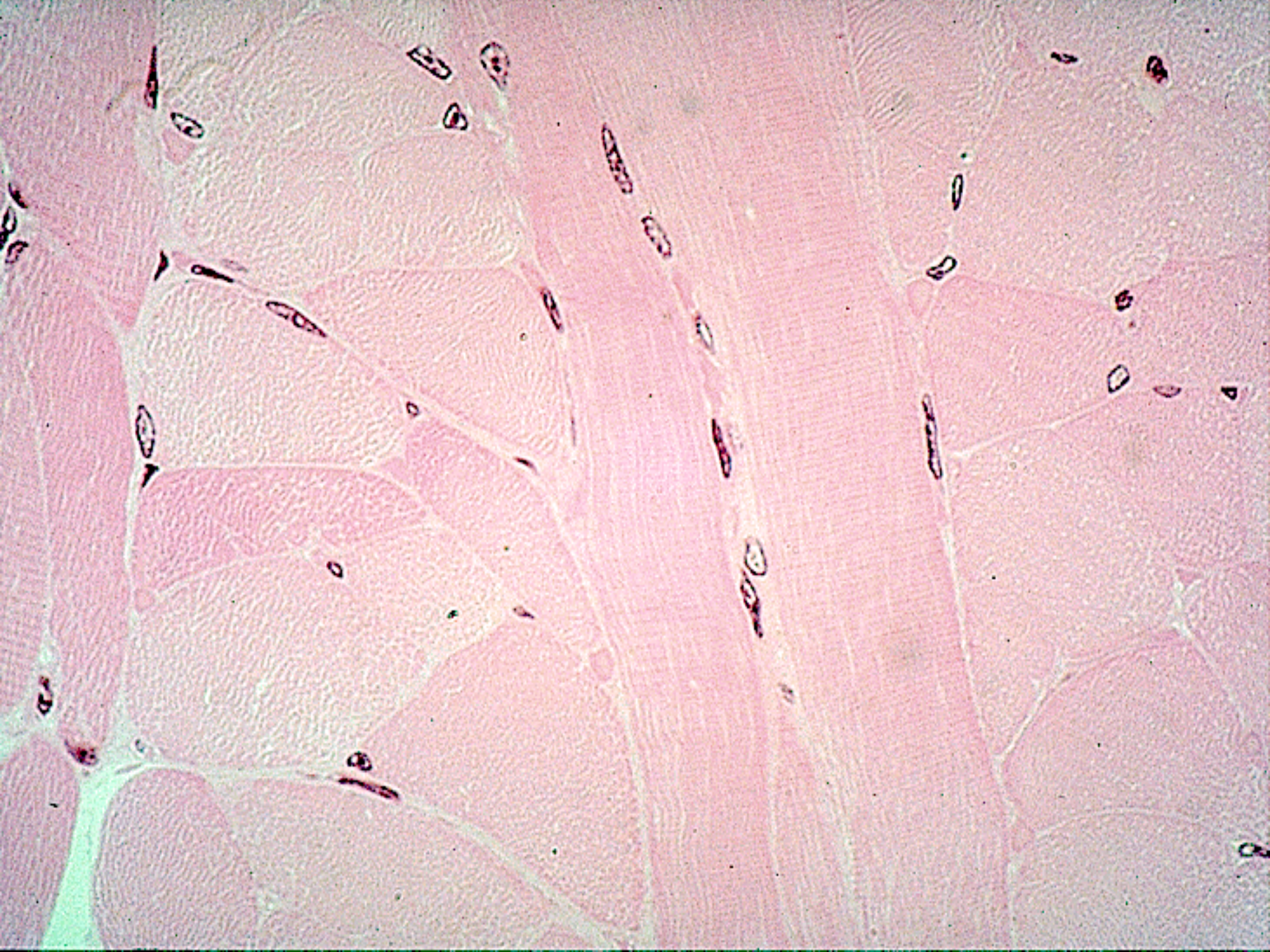
serosa

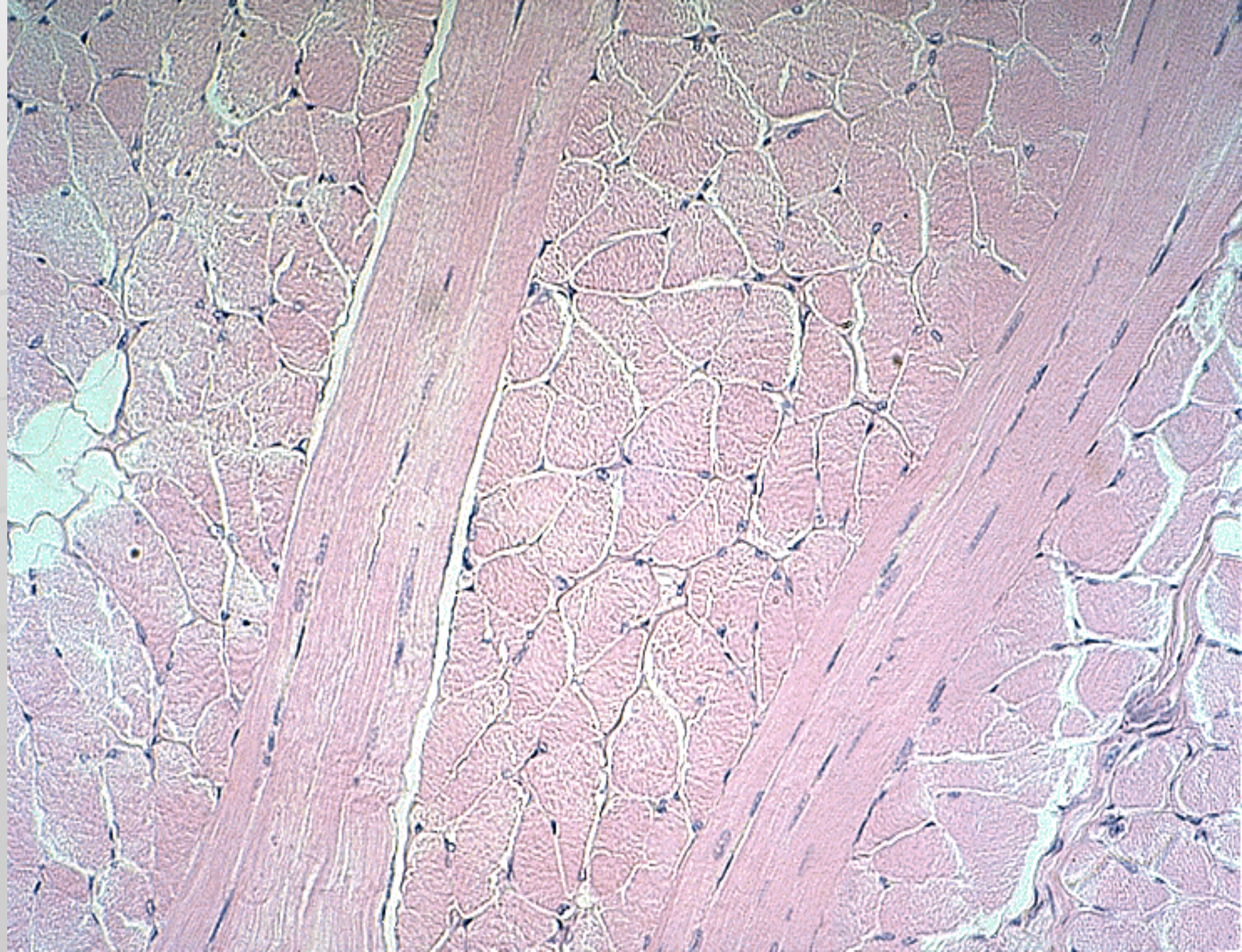
mesothelium

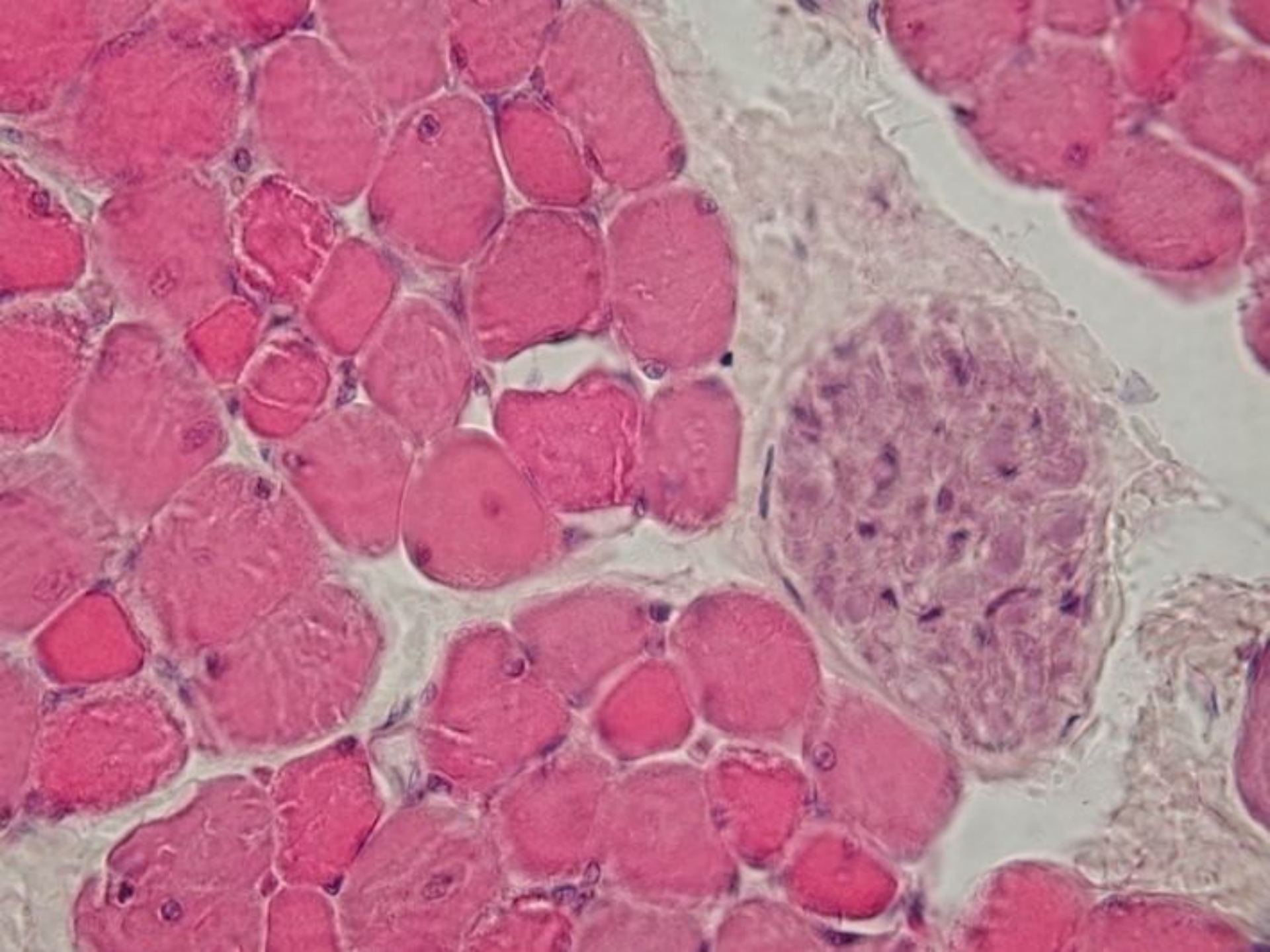


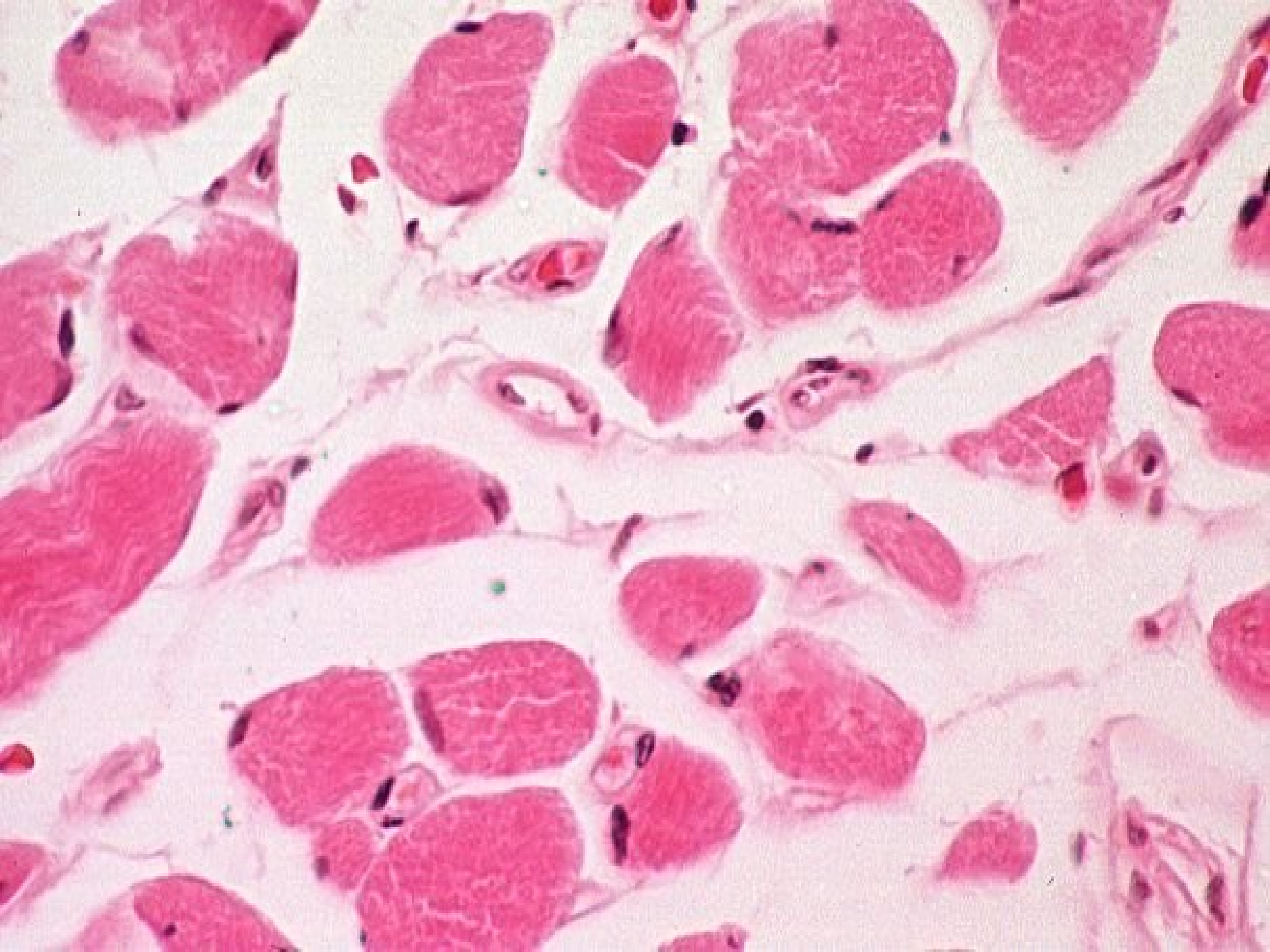
collagen

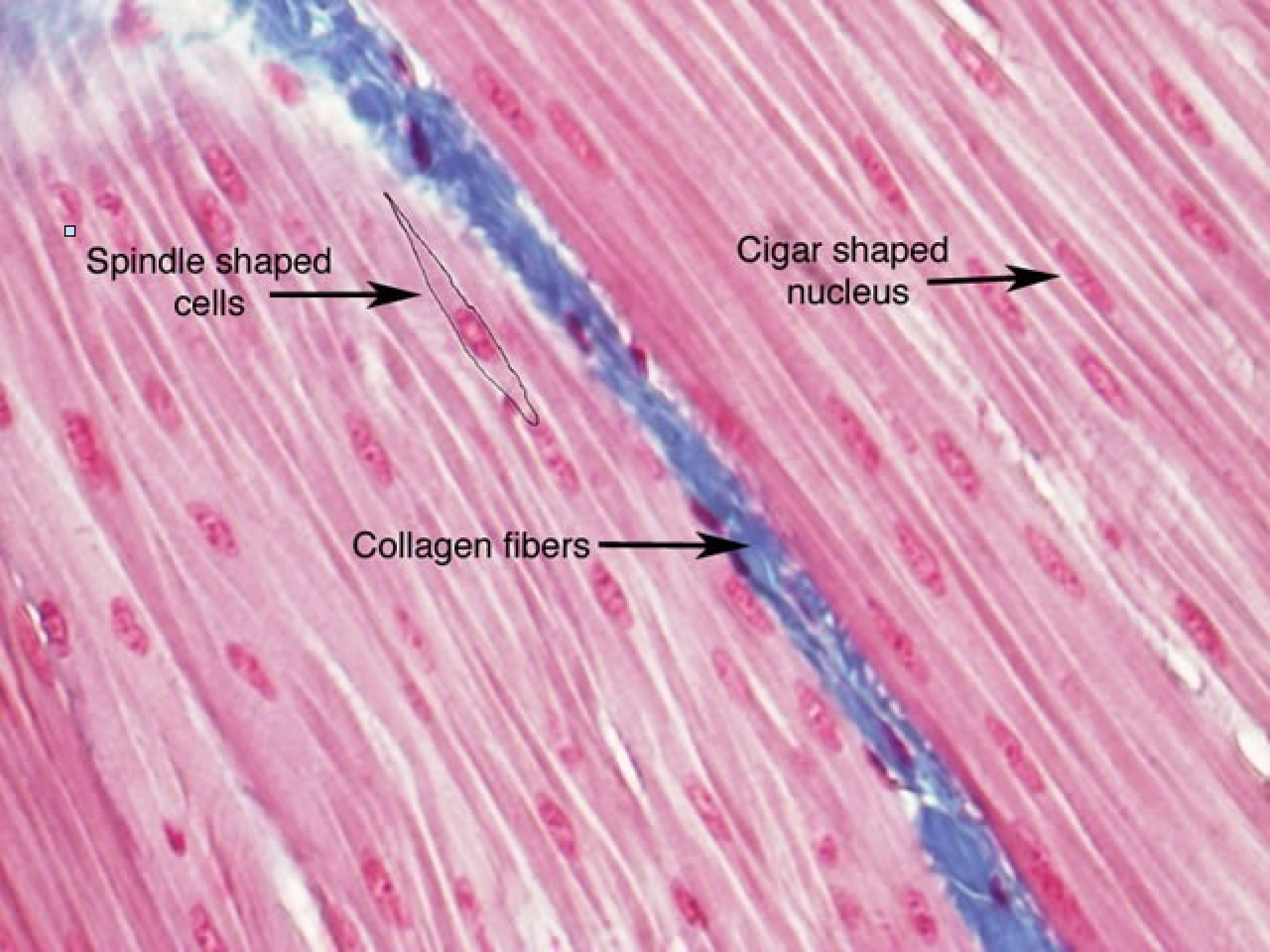










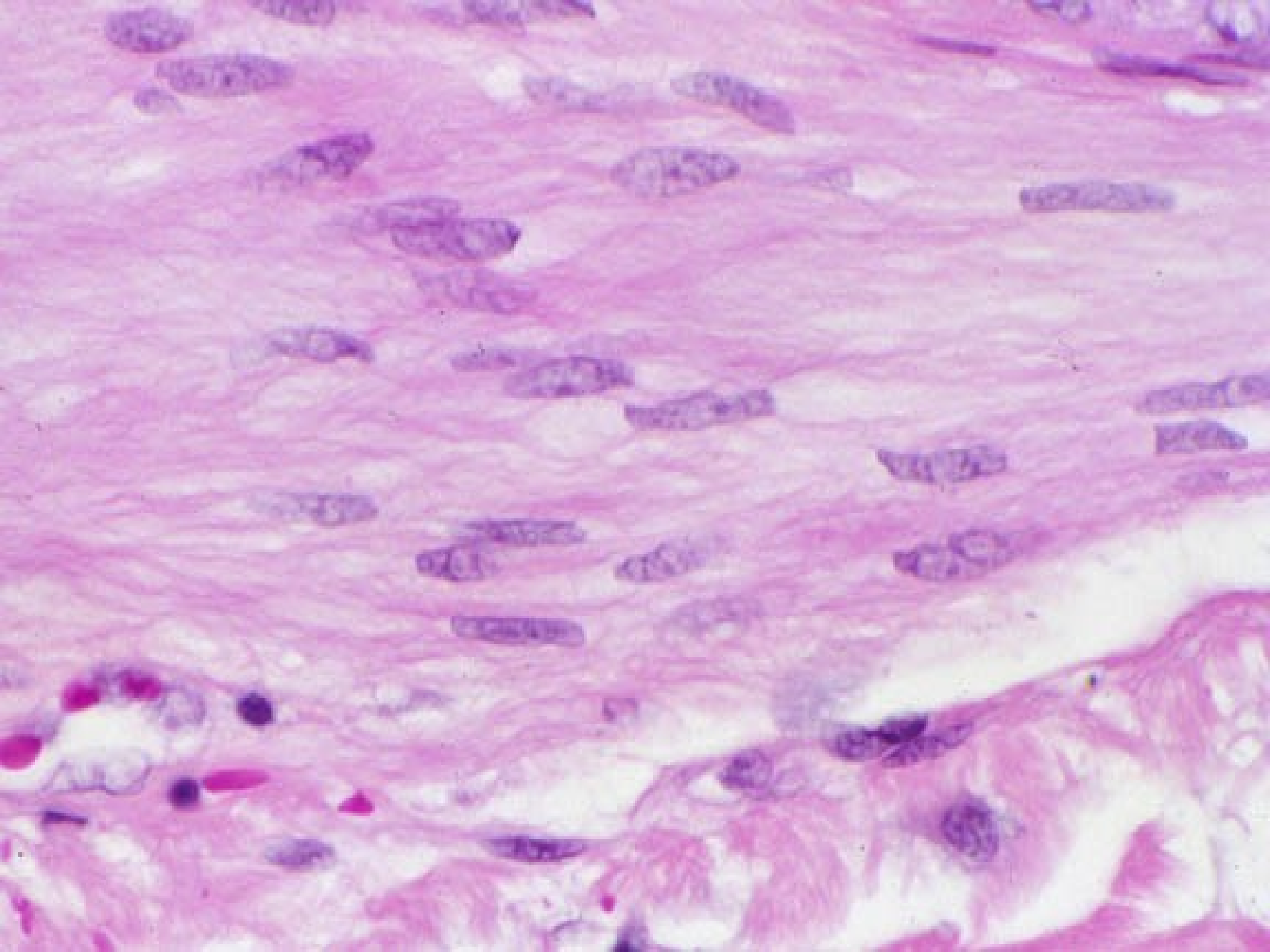


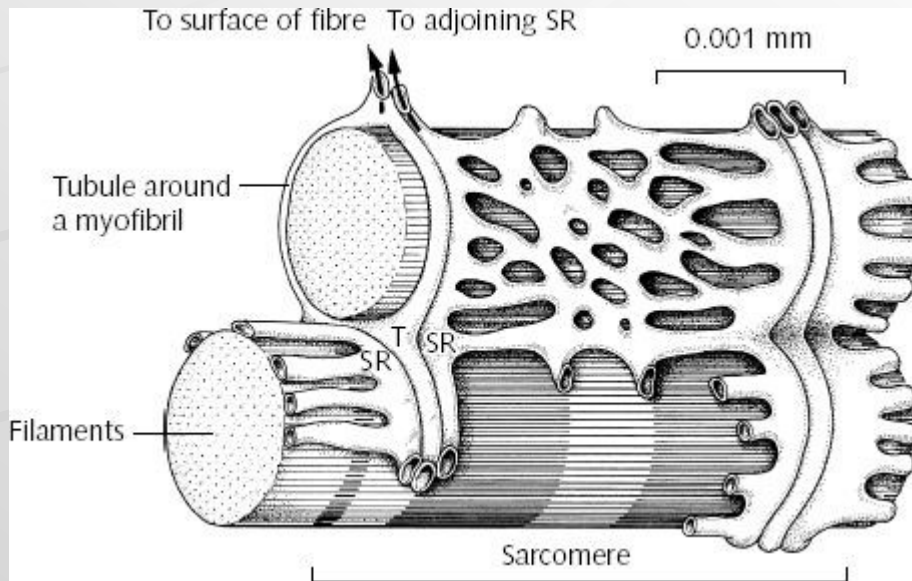
□

Spindle shaped cells

Cigar shaped nucleus

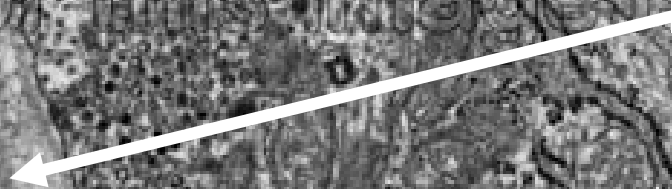
Collagen fibers





**Smooth
muscle cell**

Basal lamina



© & Photo: H. Jastrow

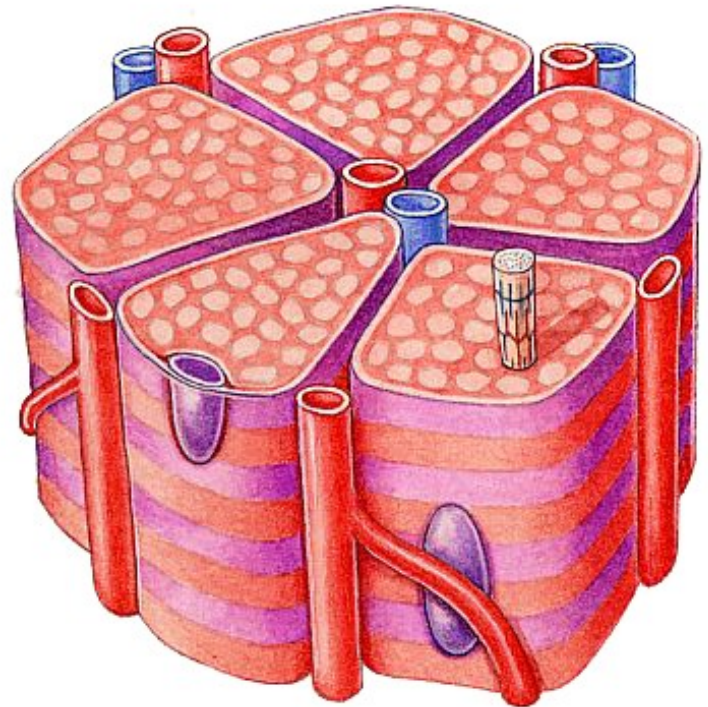
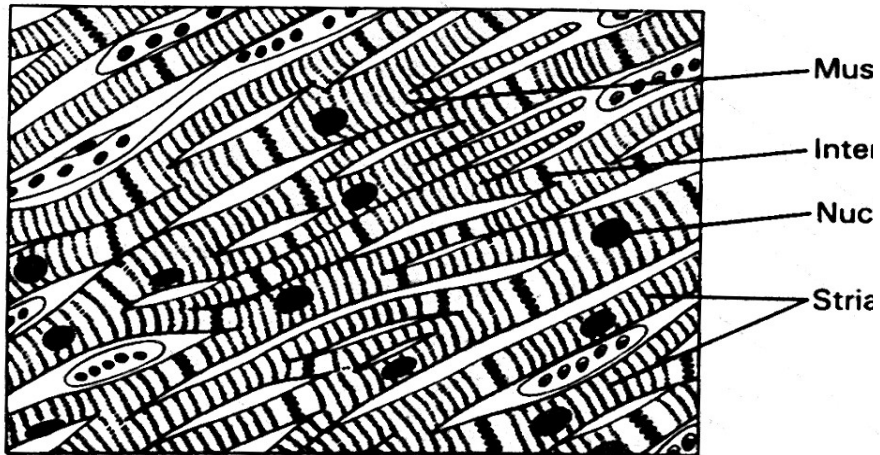
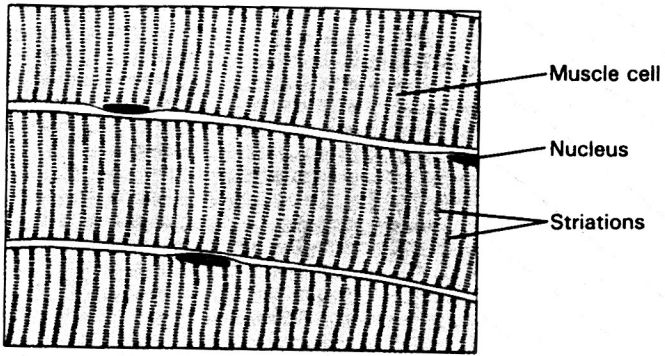
For a licensed version send an E-mail to

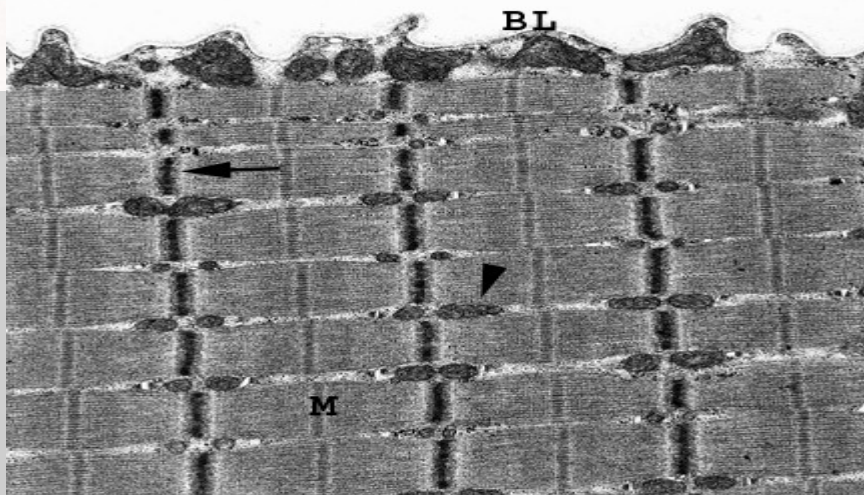
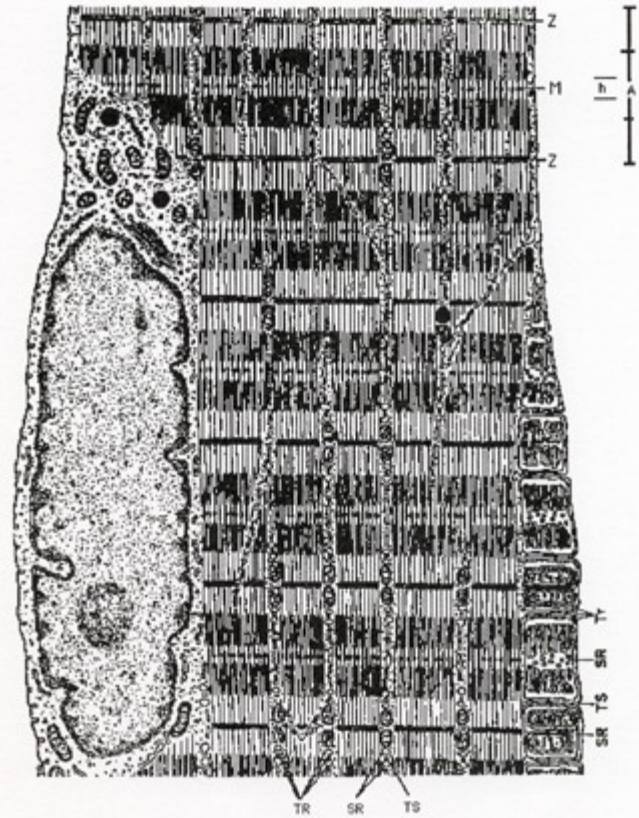
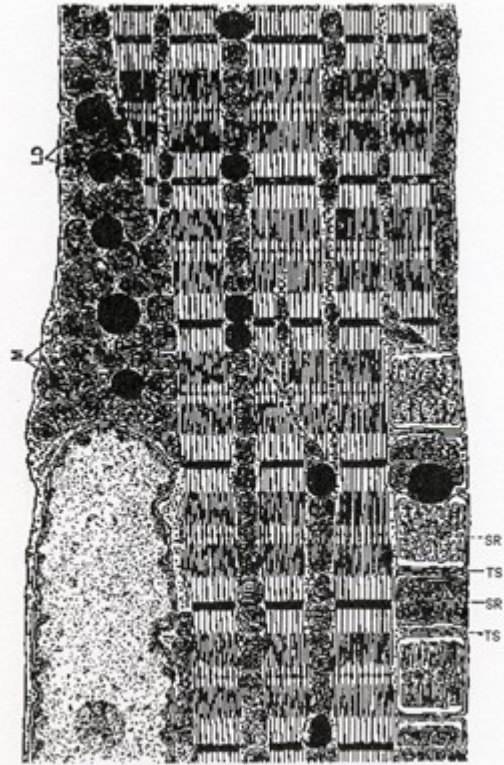
info@drjastrow.de, please

250 nm

Copyright H. Jastrow

???





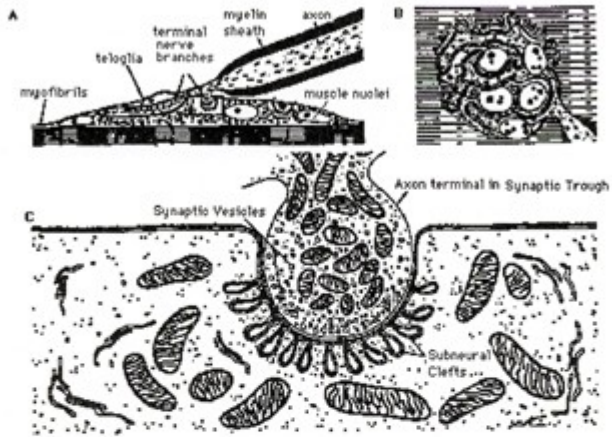
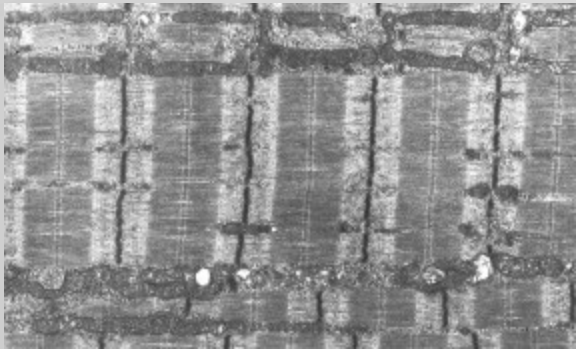


FIG. 10-9 SCHEMATIC REPRESENTATIONS OF MOTOR END PLATE AS SEEN BY LIGHT AND ELECTRON MICROSCOPY.

A. End plate as seen in histological sections
in the long axis of the muscle fiber

B. As seen in surface view with the light microscope

C. As seen in electron micrograph of an area of such
as that in the rectangle on "A".



Structure or rhabdomyocyte

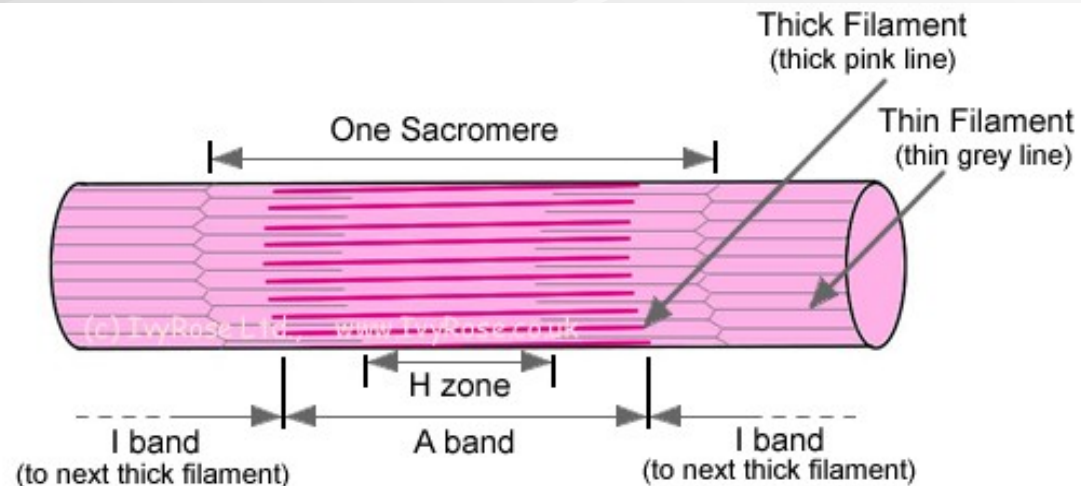
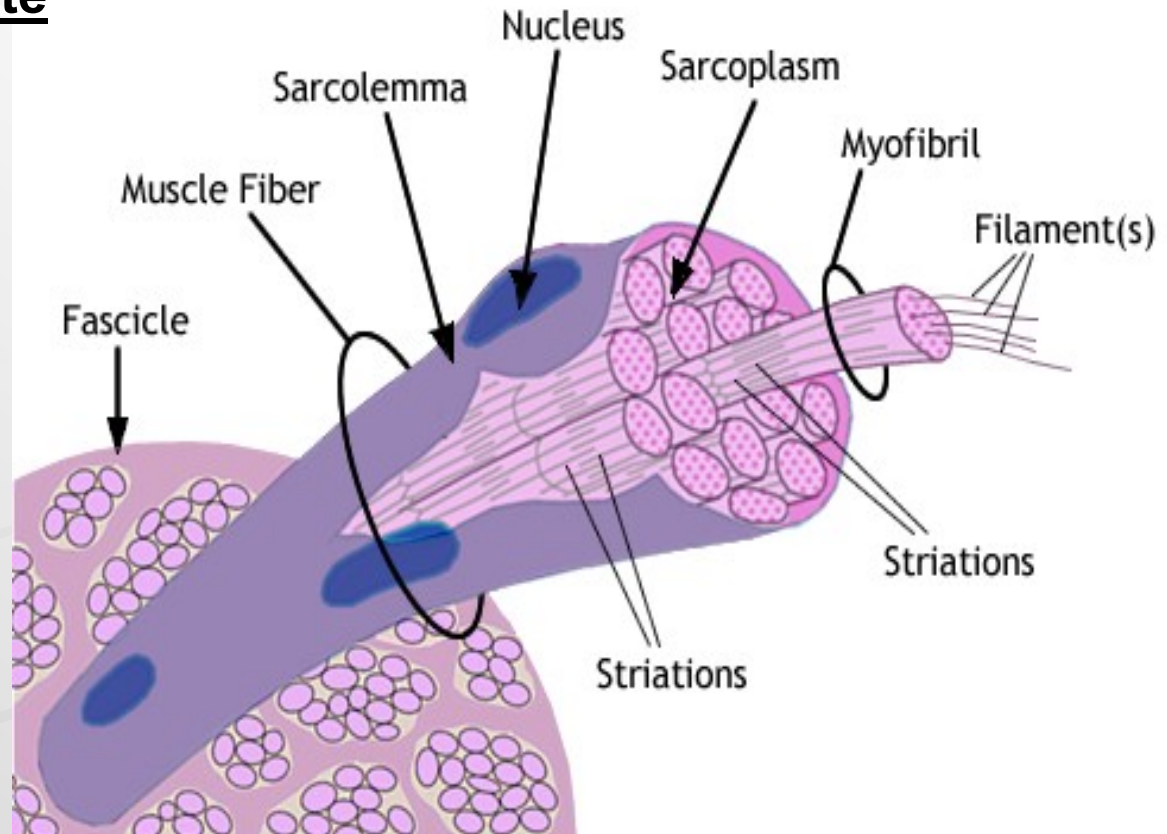
Sarcolemme + t-tubules,

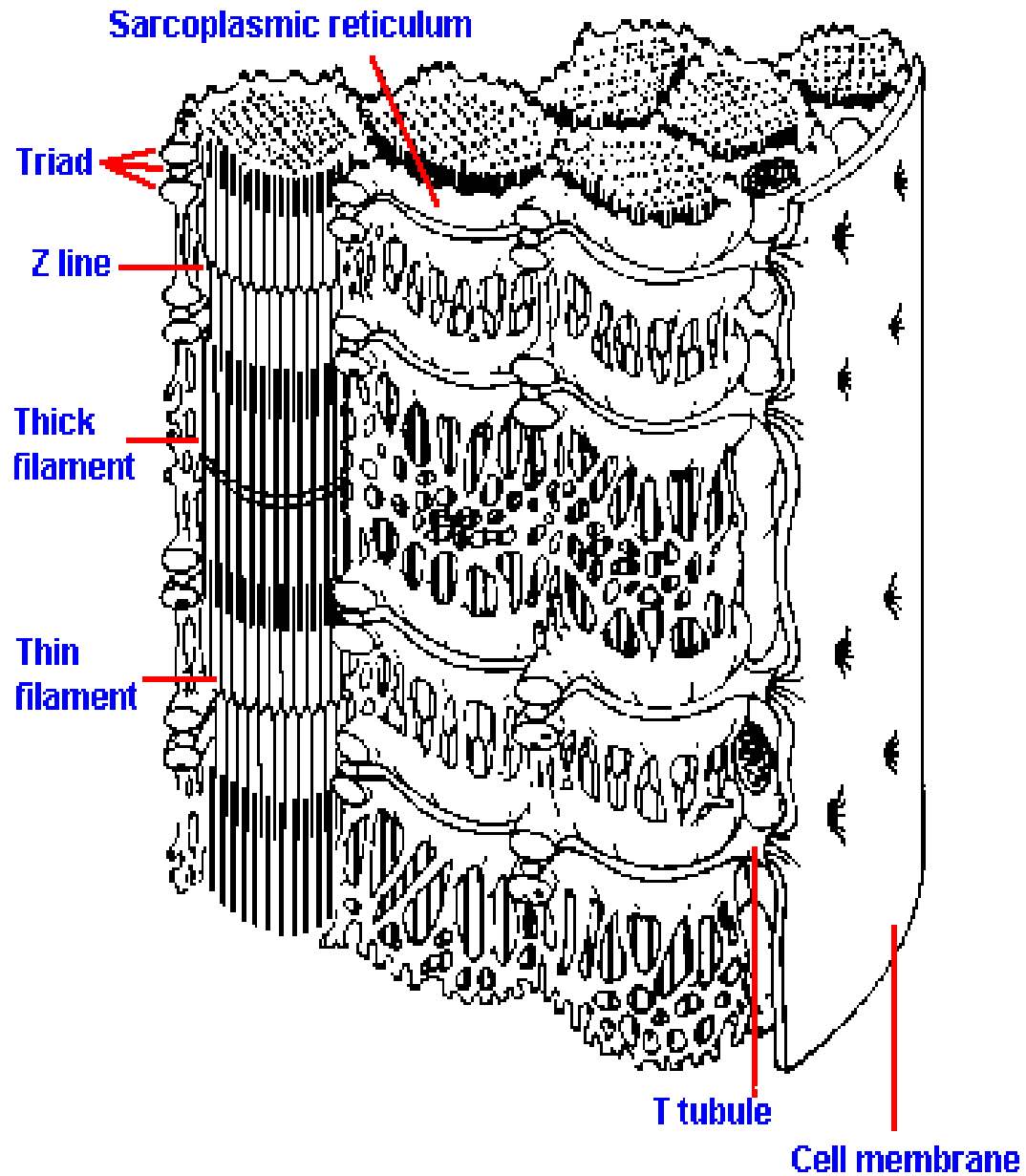
In **sarcoplasm**:

Nuclei,
Mitochondria,
Golgi apparatus,
Glycogen (beta granules)
(*sarcoplasm with organelles forms columns among myofibrils*)

Sarcoplasmic reticulum
(smooth ER) – reservoir of Ca^{2+}

Myofibrils (parallel to the length of the muscle fiber)







Mechanism of contraction:

The chemical players in muscle contraction are:

- ✚ 1. myosin (protein)
- ✚ 2. actin (protein)
- ✚ 3. tropomyosin (protein)
- ✚ 4. troponin (protein)
- ✚ 5. ATP (nucleotide)
- ✚ 6. calcium ions

- ✚ - There are two binding sites on each myosin head, one for ATP and one for actin. ATP is hydrolyzed into ADP + phosphate and energy is transferred into the head of myosin.
- ✚ - Thin filaments are made of these three protein molecules: 1. actin, 2. tropomyosin,
- ✚ 3. troponin.
- The major component of the thin filament, actin is composed of a double strand of
- ✚ actin subunits each of which contain myosin binding sites.
- The regulatory protein tropomyosin twists around the actin. When the sarcomere
- ✚ is not shortening, the position of the tropomyosin covers the binding sites on the
- ✚ actin subunits and prevents myosin cross bridge binding.
- Troponin, which is found periodically along the tropomyosin strand, functions to
- ✚ move the tropomyosin aside, exposing the myosin binding sites.
- Role of calcium in muscle contraction: action potential occurs - calcium ions are
- ✚ released from the terminal cisternae - calcium ions then bind to troponin -
- ✚ tropomyosin moves away from the myosin binding sites on actin.

Review of participants in the Cross Bridge Cycle:

<u>Participant</u>	<u>Will bind to:</u>
1. Myosin	ATP, Actin
2. Actin	Myosin, Troponin
3. Tropomyosin	Troponin
4. Troponin	Calcium, Actin, Tropomyosin
5. ATP	Myosin
6. Calcium ions	Troponin

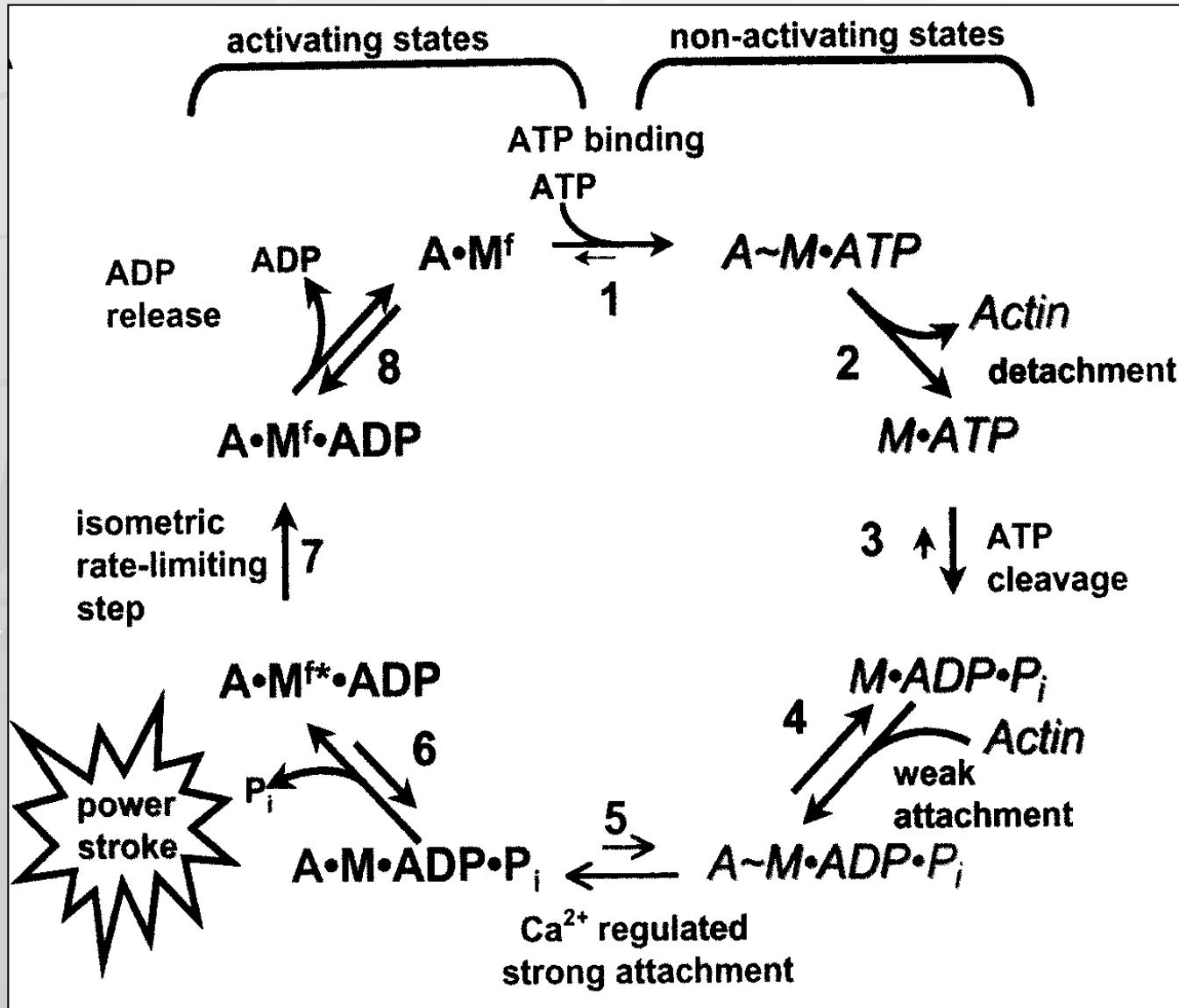
Detail of steps required to expose the binding sites on actin:

- ✦ Presence of an action potential in the muscle cell membrane.
- ✦ Release of calcium ions from the terminal cisternae.
- ✦ Calcium ions rush into the cytosol and bind to the troponin.
- ✦ There is a change in the conformation of the troponin-tropomyosin complex.
- ✦ This tropomyosin slides over, exposing the binding sites on actin.

Summary of the role that ATP plays in the contraction of muscle:

- ➡ 1. ATP transfers its energy to the myosin cross bridge, which in turn energizes the power stroke.
- ➡ 2. ATP disconnects the myosin cross bridge from the binding site on actin.
- ➡ 3. ATP fuels the pump that actively transports calcium ions back into the sarcoplasmic reticulum.

ATP is required to cleave the avidly formed bond between actin and myosin. In the absence of ATP the cross linking becomes fixed, and the muscle becomes rigid and inextensible. This condition is seen shortly after death when the muscles' energy supply is used up, and is known as **rigor mortis**.



The thin actin filament is a dimeric polymer of G-actin sub-units arranged like two strings of beads twisted together.

Attached to the actin chain of the thin filament, are the proteins **troponin** (Tn) and **tropomyosin**.

A **tropomyosin** molecule runs along each actin chain, bound to the actin. Each tropomyosin sub-unit covers about 7 G-actin sub-units.

The **troponin** molecule has three sub-units: **TnT** that binds to tropomyosin near the ends of the tropomyosin sub-units; **TnI** that binds to the actin; and **TnC** that binds to the TnI and TnT sub-units, and which also has a strong affinity for Ca^{2+} at four binding sites.



The thick filaments, at either end, have sticking off at regular intervals, the fibrillar necks and globular, **ATPase** heads of the myosin subunits - the polymerized tails of which make the backbone of the filament.

The heads at the opposite ends of the thick filament stick off in opposite directions leaving a bare middle region (pseudo H-zone) on the filament.

Each successive myosin head is staggered so that six rows of heads stick off in six directions (60° intervals) around the filament.

Each myosin filament is surrounded by six actin filaments and each actin filament by three myosins.



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