

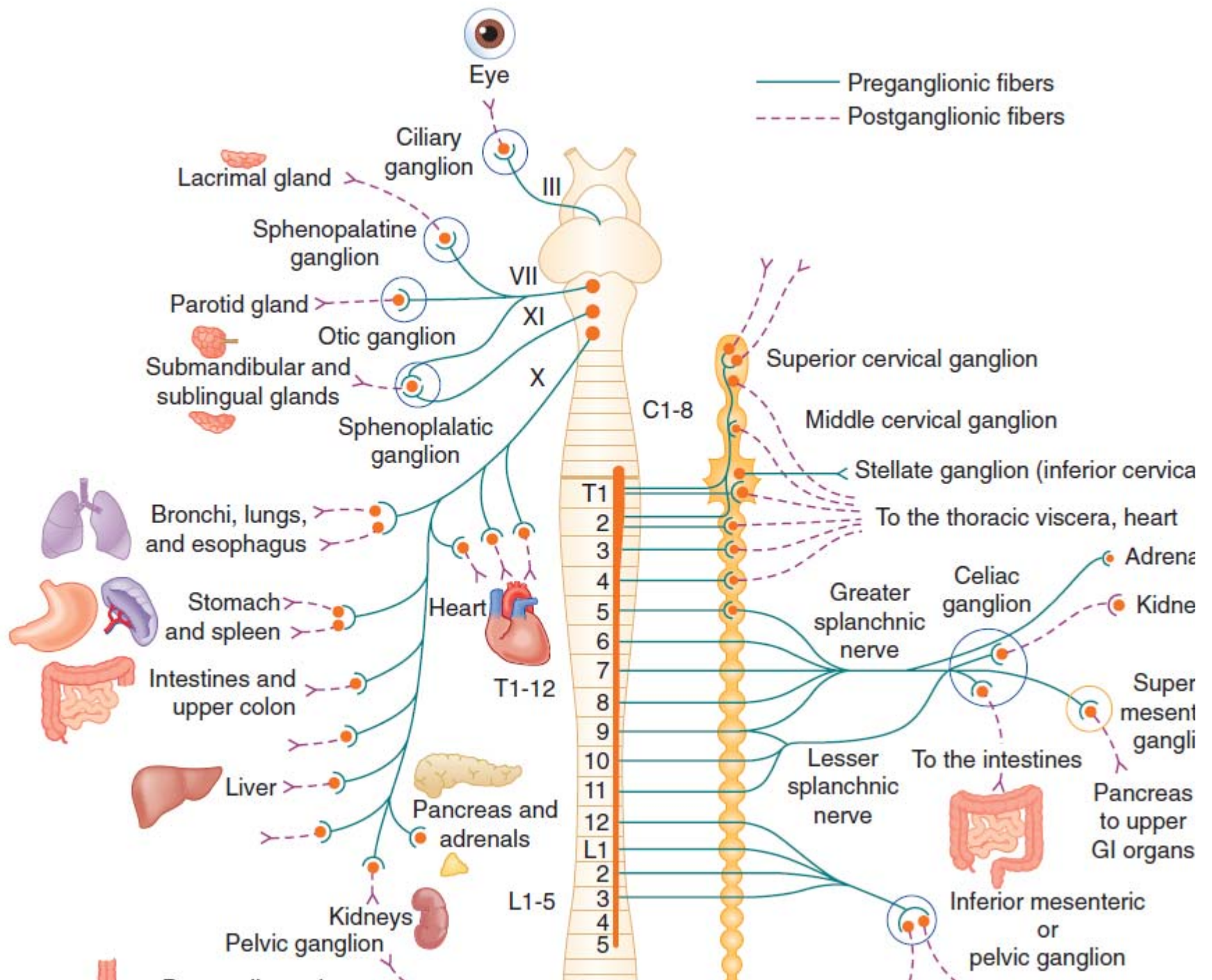
- συμπάθεια/sympathia(=>συν/syn=together, with) +πάθος / pathos =passion, sorrow, strong emotion
- Galen (Greek physician) – sympathetic nervous system coordinates body function
- Nerves were considered as hollow tubes
- Distribute animal spirit
- Foster concerned action or “syphathy”

- The term sympathetic nervous system coined by Winslow (1669-1760)
 - Described the chains of ganglia and the nerves connected to the thoracic and lumbar spinal cord
- Bichat (1771-1802)
 - divided life into two distinct forms,
 - one governed by the brain,
 - and the other (organic, vegetative, life) by the abdominal ganglia → Little Brains

- English physiologist John Langley (1852-1925)

introduced the term autonomous nervous system and parasympathetic nervous system
-the cranial and sacral portions of the nervous system

- Παρά/para(= prefix denoting identification, “beside”, “near”, “contrary of” “side by side”



AUTONOMIC NERVOUS SYSTEM

PEIPHERAL PART

Three major divisions:

- SYMPATHETIC
- PARASYMPATHETIC AND
- ENTERIC.

CENTRAL AUTONOMIC NETWORK

- SPINE
- LOWER BRAINSTEM (BULBO-PONTINE)
 - NTS
 - RVLM
 - Parabrachial Nuclei

Reflex control of circulation, Respiration, GIT functions and Micturition
- UPPER BRAINSTEM(Ponto Mesencephalic) – Periaqueductal grey
 - Integrate autonomic control with pain
 - Behavioral response to stress and Sleep
- FOREBRAIN – Hypothalamus
- Anterior Limbic System – Insula, Anterior Cingulate gyrus and Amygdala
 - Integration of bodily sensation with emotional and goal related autonomic response

FUNDAMENTAL FEATURE

- **Convergence of**

- Visceral afferents
- Nociceptive Inputs
- Thermoceptive and
- From muscle Receptors

All these signals inform the CAN about the physiological state of the body

Integrated and then Parallel information is send to Parasympathetic and Sympathetic system

FUNDAMENTAL DIFFERENTIAL CRITERIA FOR PARASYMPATHETIC AND SYMPATHETIC NERVOUS SYSTEM

- **Segmental Organization of their Preganglionic Neurons**
- **The Peripheral location of the ganglia**
- **The type and Location of their end organ**
- **The effect they produce on end organs**
- **The neurotransmitters employed by their Post ganglionic neurons**

The sympathetic and parasympathetic divisions innervate

- cardiac muscle, smooth muscle, and glandular tissues and
- mediate a variety of visceral reflexes

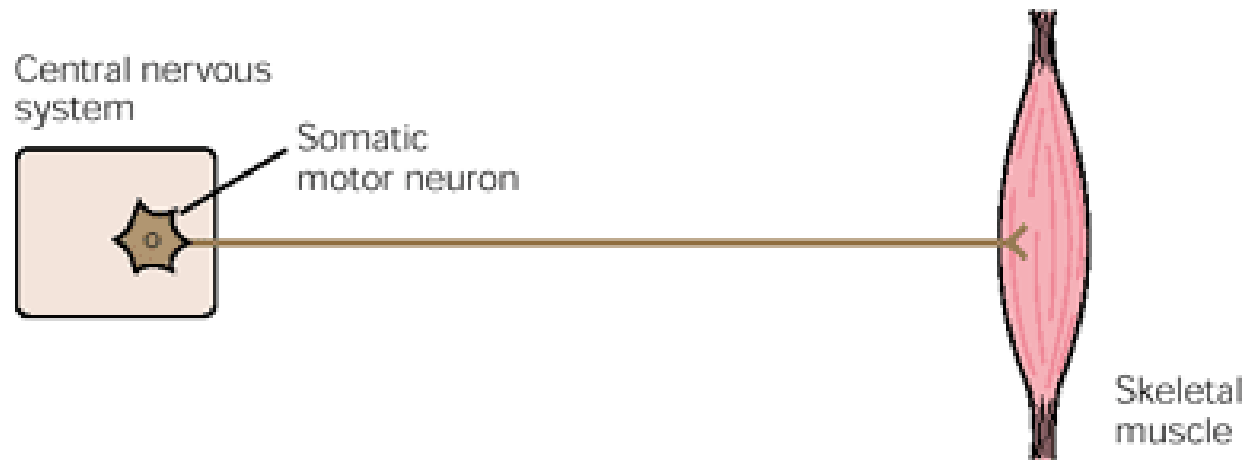
These two divisions include

- the sensory neurons associated with spinal and cranial nerves
- the preganglionic and postganglionic motor neurons
- the central nervous system circuitry that connects with and modulates the sensory and motor neurons.

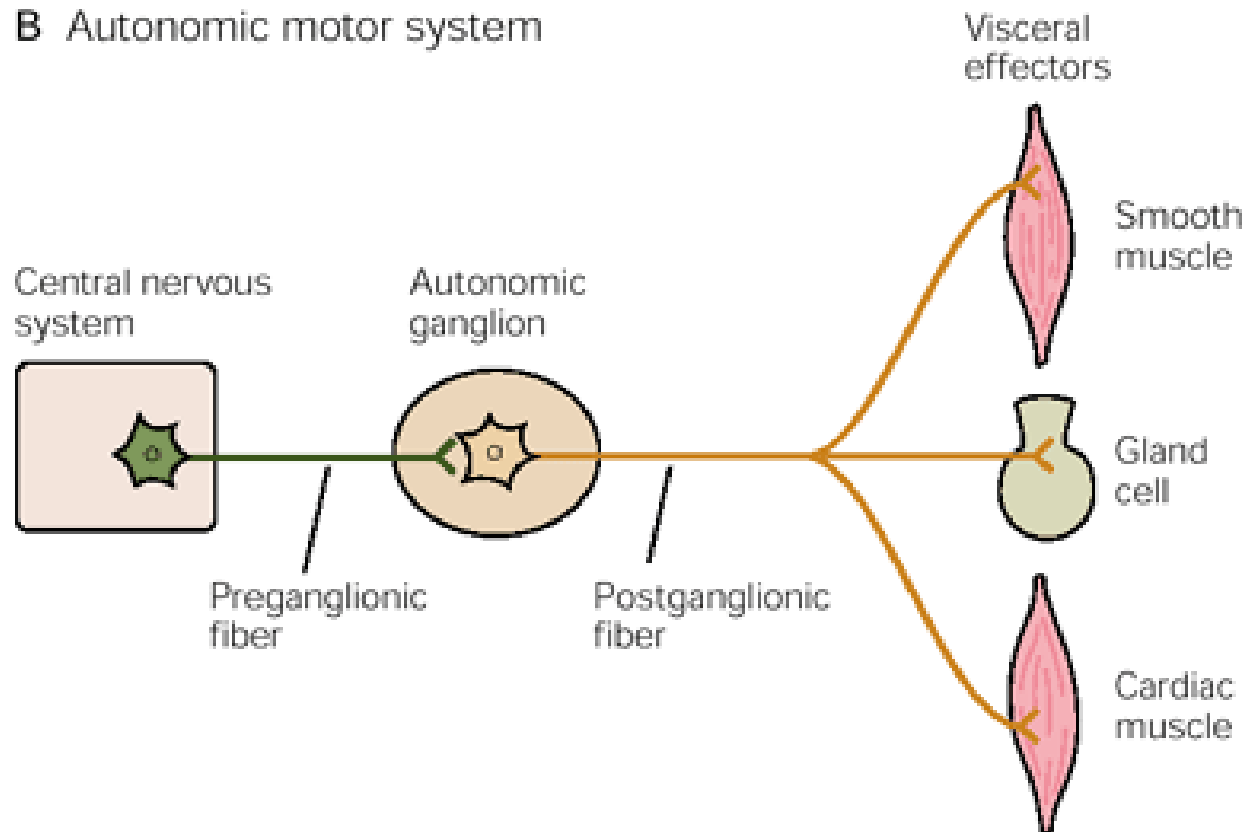
- *The enteric division has greater autonomy*
 - comprises a self-contained system, with only minimal connections to the rest of the nervous system.
 - It consists of sensory and motor neurons in the gastrointestinal tract that mediate digestive reflexes.
- *Parasympathetic nervous system*
 - responsible for *rest and digest*
 - *maintain basal heart rate, respiration, and metabolism under normal conditions.*

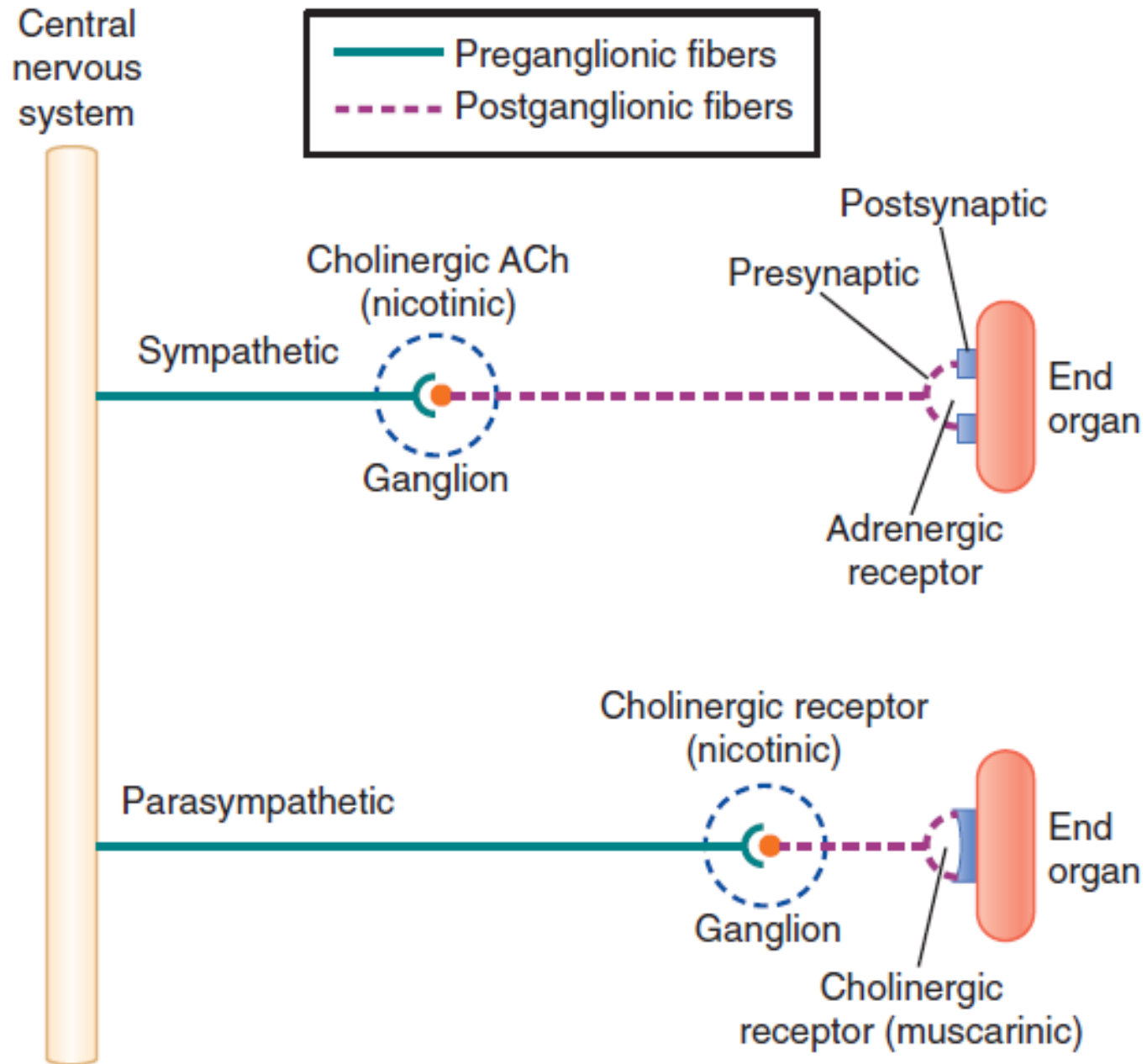
-
- *The sympathetic nervous system*
 - governs the emergency reaction, or *fight-or-flight reaction*.
- For a person to respond effectively
the sympathetic nervous system increases
 - output to the heart and other viscera,
 - the peripheral vasculature and sweat glands,
 - and the piloerector and certain ocular muscles.

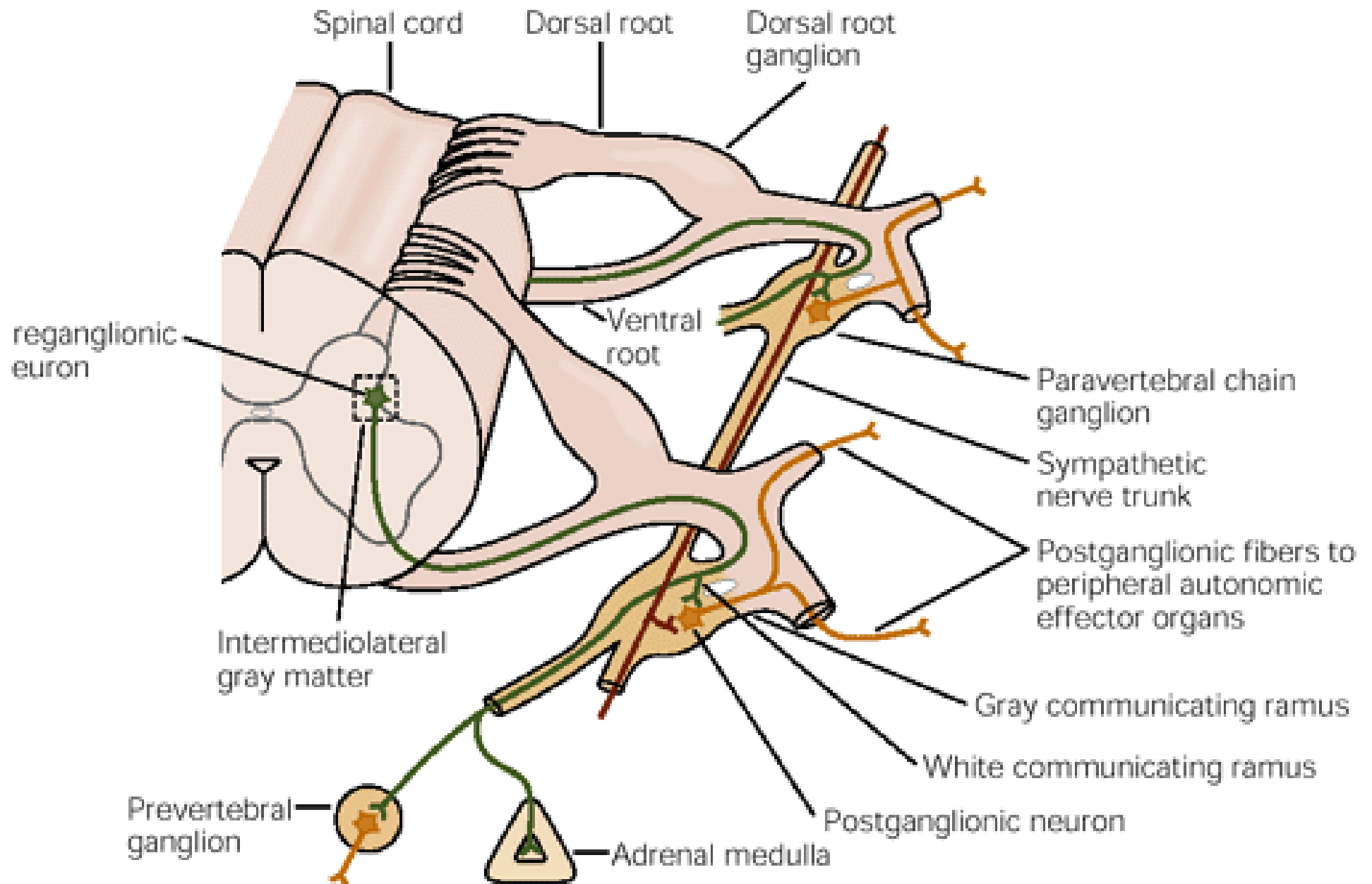
A Somatic motor system



B Autonomic motor system







SYMPATHETIC NERVOUS SYSTEM

- 22 pairs of **Paravertebral ganglia**
- Ganglia connected to each other via nerve trunk
- Ganglia connected to Spinal nerves by *rami communicantes*
- *White rami* – from each thoracolumbar segment, carry preganglionic myelinated fibres to ganglia
- *Grey rami* – carry postganglionic unmyelinated fibres back to the spinal nerves to be distributed to
 - --Sweat glands
 - --*Pilomotor muscles*
 - --*Blood vessels of skeletal muscle and skin*
- Follows the course of spinal nerves or blood vessels or both

- Prevertebral ganglia
 - In abdomen and Pelvis
 - Celiac, Superior mesenteric, Aorticorenal and Inferior mesenteric
- Terminal ganglia
 - Lie near the organs- Urinary bladder, Rectum and
 - Cervical ganglia in the neck
 - Many sympathetic preganglionic fibers from 5th to last thoracic segment pass through paravertebral ganglia –form Splanchnic nerves– Reach celiac ganglia and then synapse
 - Some pass directly to the Adrenal medulla
 - Post ganglionic fibers secrete epinephrine

Segments	Output Ganglia	Effectors	Effect (Neurotransmitter)
T1–T3	Superior cervical	Eye Salivary glands Face sweat glands Blood vessels Pineal gland	Pupil dilation (NE) Contraction of tarsal muscle (NE) Increase in salivary protein (NE) Facial sweating (ACh) Vasoconstriction (NE) Melatonin secretion (NE)
T1–T8	Stellate and upper thoracic (upper limb, and thoracic viscera)	Sweat glands Skin blood vessels Piloerector muscles Muscle blood vessels Periosteum, bone	Sweating (ACh) Skin vasoconstriction (NE) or vasodilation (NO) Piloerection Muscle vasoconstriction (NE) or vasodilation Bone marrow activation (NE)
T5–T12	Celiac Mesenteric Adrenal medulla	Heart Respiratory tract Abdominal viscera Blood vessels Endocrine cells	Cardiac stimulation (NE) Bronchodilation (NE) Inhibition of GI motility Vasoconstriction Renin secretion Glucagon secretion Epinephrine secretion
T10–L2	Lumbar paravertebral	Sweat glands Skin blood vessels Piloerector muscles Muscle blood vessels	Sweating (ACh) Skin vasoconstriction (NE) or vasodilation (NO?) Piloerection Muscle vasoconstriction (NE) or vasodilation
T2–L2	Inferior mesenteric Hypogastric	Pelvic viscera and blood vessels	Urine storage Rectal inhibition Ejaculation

- Post ganglionic fibers supply visceral structures of thorax, abdomen, head and neck.
- The trunk and limbs get their sympathetic supply in spinal nerves
- Many of upper thoracic sympathetic fibers from vertebral ganglia form terminal plexuses like Cardiac, Esophageal and Pulmonary
- Head and Neck – Cervical sympathetic chain and its three ganglia
 - All preganglionic fibers arise from upper thoracic segments

Parasympathetic System

- Cranial component
 - Supply Sphincter of Iris
 - Ciliary muscle
 - Salivary glands
 - Lacrimal glands
 - Mucous glands of nose , mouth and pharynx
- The Vagus nerve
 - supply viscera of thorax and abdomen
- Parasympathetic Sacral Outflow
 - Pelvic nerve → Bladder, Rectum and Sex organs

- Axons of preganglionic neurons exit the spinal cord at the level at which their cell bodies are located,
- → *they may innervate sympathetic ganglia situated either above or below in the sympathetic nerve trunk that connects the ganglia*
- **Most of the preganglionic axons are relatively slow-conducting, small-diameter myelinated fibers.**
- Each preganglionic fiber forms synapses with many postganglionic neurons in different ganglia, the ratio
- 1:10 → Divergence → coordinated activity in sympathetic neurons at several different spinal levels

- *The axons of postganglionic neurons - unmyelinated* → Exit the ganglia in the *gray rami*.
- *The postganglionic cells innervating the head* are located in the superior cervical ganglion
- *The axons of these cells travel along branches of the carotid arteries to their targets in the head.*
- The postganglionic fibers innervating the rest of the body travel in spinal nerves to their targets;
- In an average spinal nerve, about 8% *of the fibers are sympathetic postganglionic axons.*

- *Some neurons of the cervical and upper thoracic ganglia innervate cranial blood vessels, sweat glands, and hair follicles*
- others innervate the glands and visceral organs of the head and chest, lacrimal and salivary glands, heart, lungs, and blood vessels
- *Neurons in the lower thoracic and lumbar paravertebral ganglia innervate peripheral blood vessels, sweat glands, and pilomotor smooth muscle*

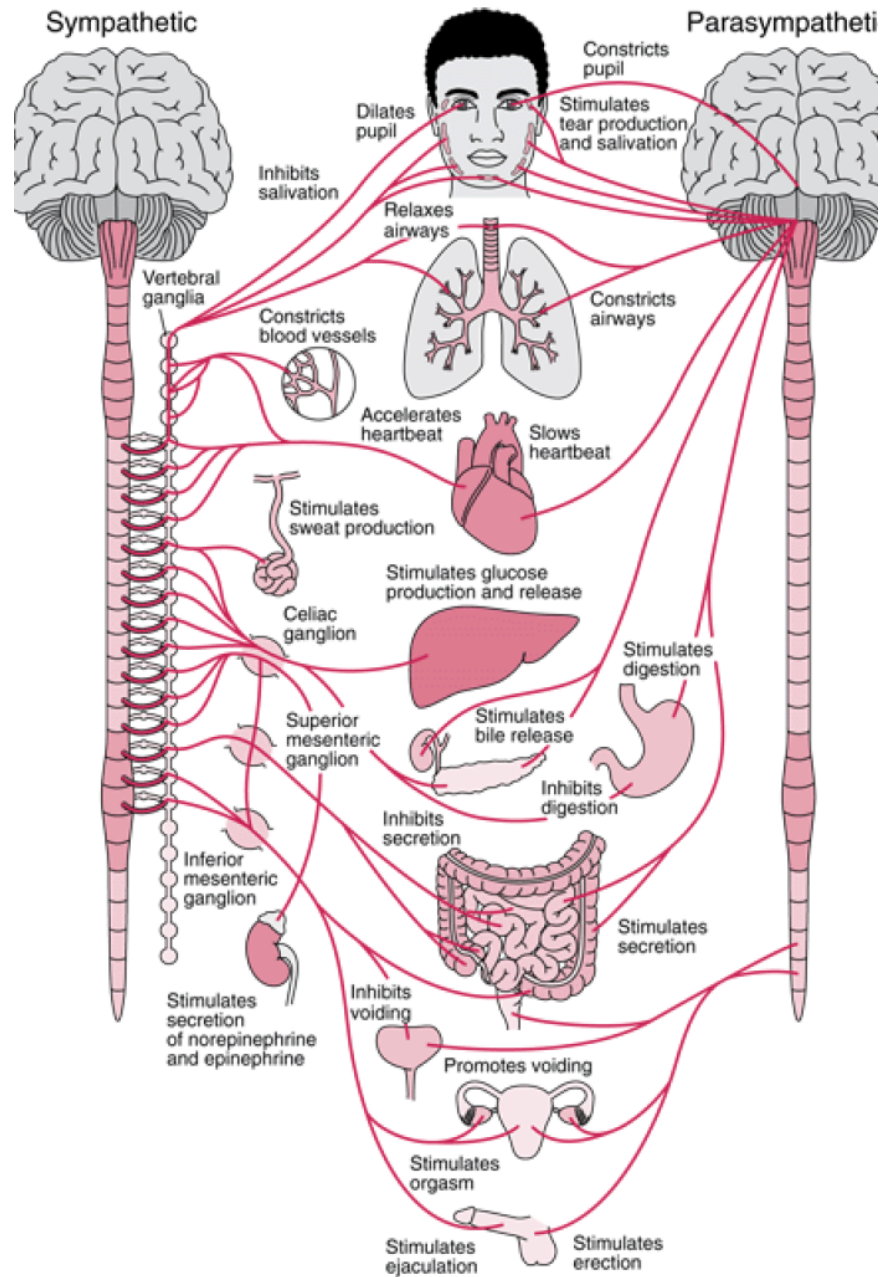
- Some preganglionic fibers pass through the sympathetic ganglia and branches of the splanchnic nerves to synapse on neurons of the *prevertebral ganglia*- the coeliac ganglion and the superior and inferior mesenteric ganglia
 - *supply whole of the GIT and kidneys, bladder, and genitalia.*
- Another group of preganglionic axons runs in the thoracic splanchnic nerve into the abdomen and innervates the adrenal medulla.

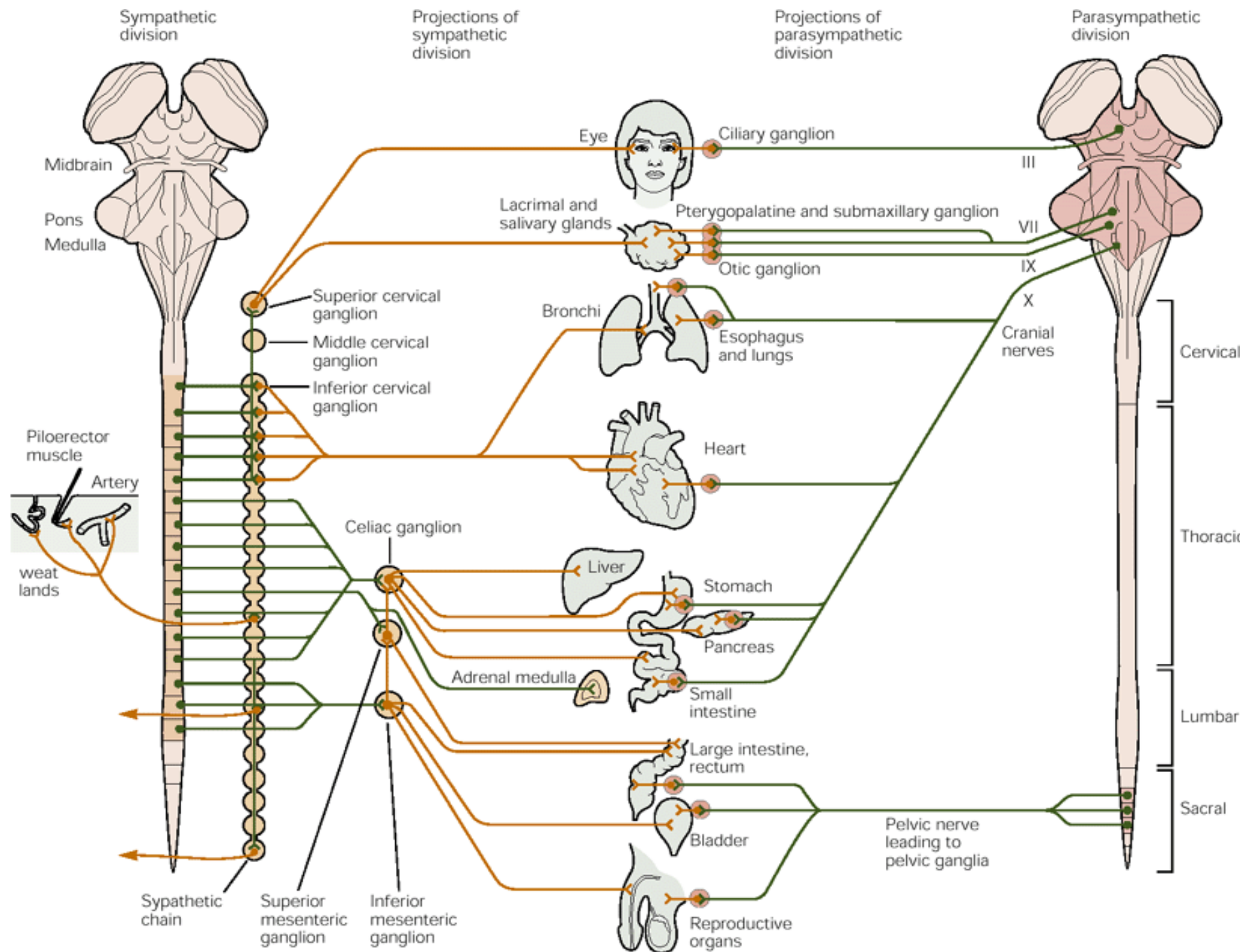
- The central, preganglionic cells of the parasympathetic nervous system are located in several brain stem nuclei and in segments S2-S4 of the sacral spinal cord
- The axons of these cells are long → parasympathetic ganglia lie close to or embedded in visceral target organs.
- The preganglionic parasympathetic nuclei in the brain stem –
 - Edinger-Westphal nucleus
 - Superior and inferior salivary Nuclei
 - Dorsal vagal nucleus and the nucleus ambiguus

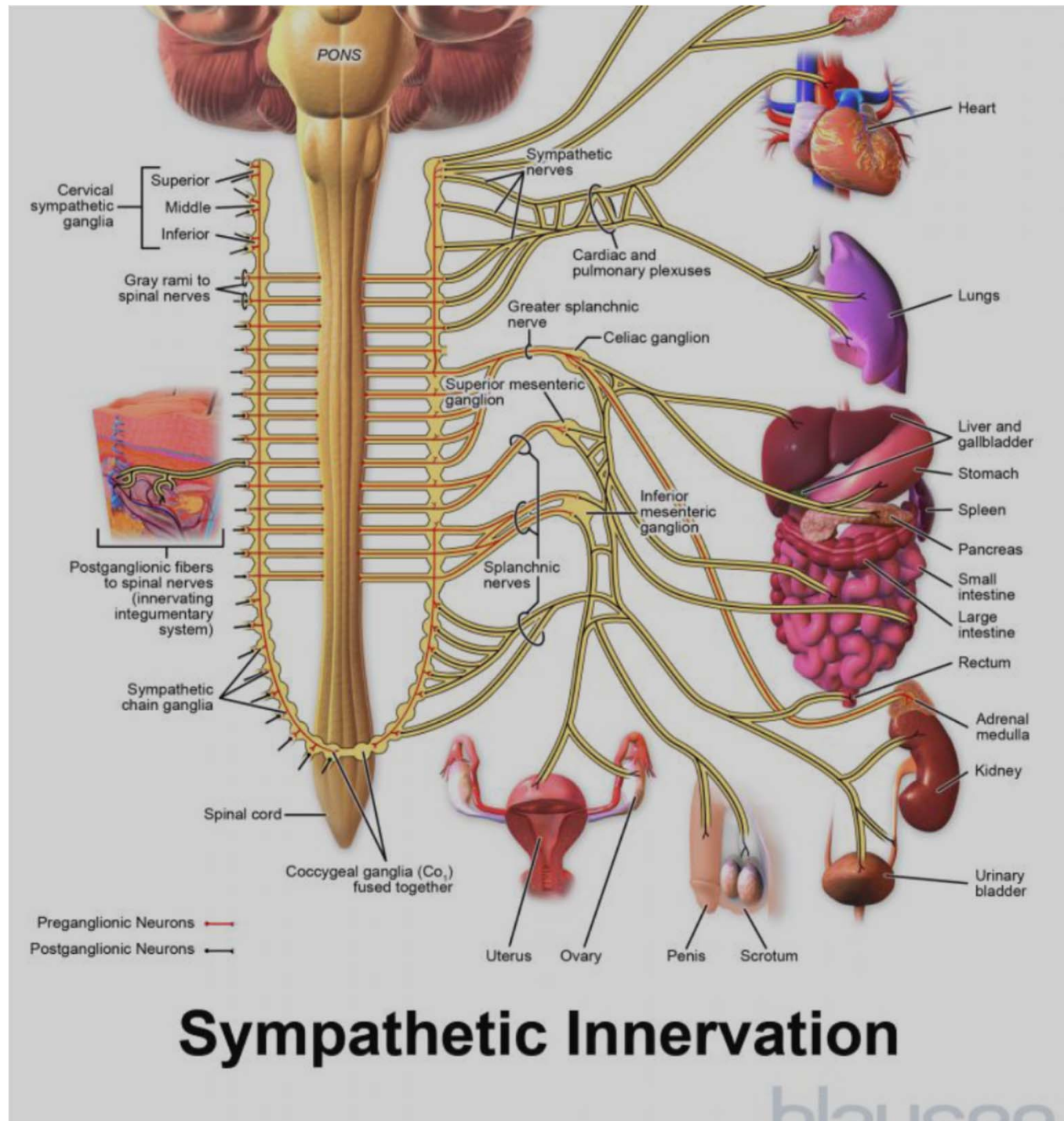
- Preganglionic axons exit the brain stem through cranial nerves III, VII, and IX → postganglionic neurons in the ciliary, pterygopalatine, submandibular, and otic ganglia
- From the dorsal vagal nucleus → nerve X to postganglionic neurons in -- stomach, liver, gall bladder, pancreas, and upper intestinal tract
- Ventrolateral nucleus ambiguus → the principal parasympathetic innervation of the cardiac ganglia- which innervate the heart, esophagus, and respiratory airways.

- In the sacral spinal cord -the parasympathetic preganglionic neurons are present in the intermediolateral column.
- Spinal parasympathetic nerves leave the spinal cord through the ventral roots → project in the pelvic nerve → pelvic ganglion plexus → innervate the descending colon, bladder, and external genitalia

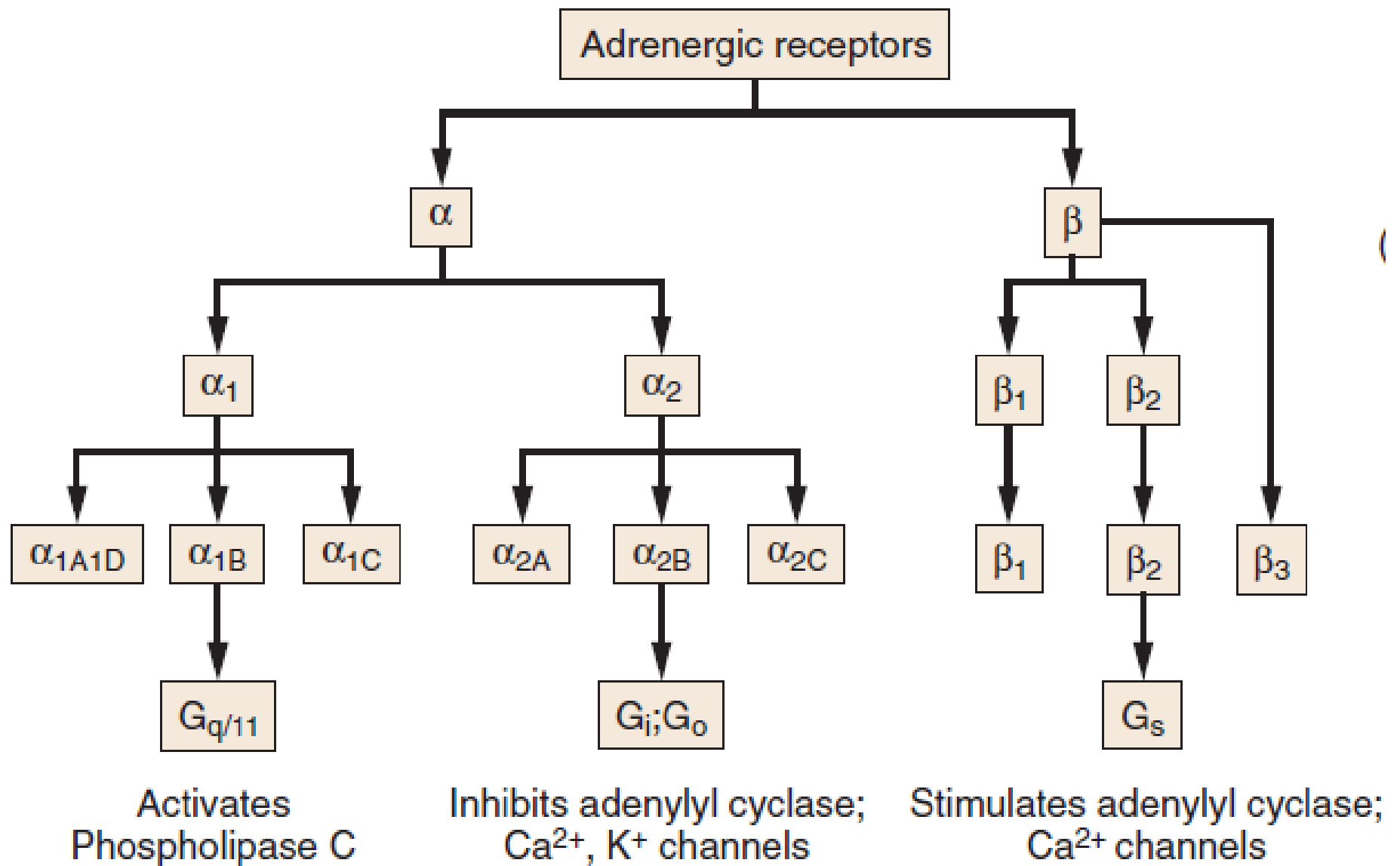
The autonomic nervous s





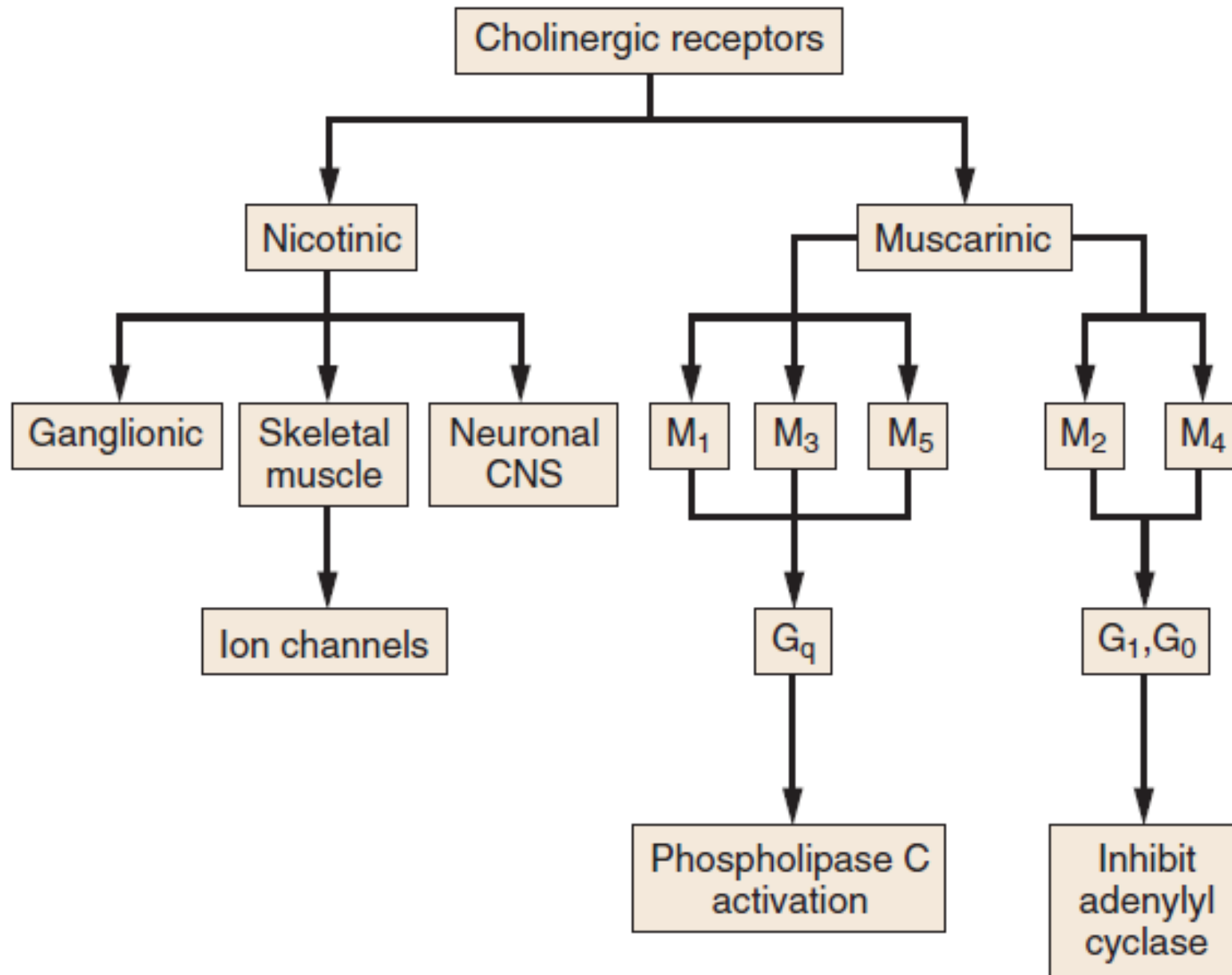


RECEPTOR	EFFECTOR	RESPONSE TO STIMULATION
Sympathetic Nervous System		
α_1	Smooth muscle (vascular, iris radial, ureter, trigone, bladder sphincters)	Constriction
α_2	Presynaptic SNS nerve endings	Inhibition of NE release
	Brain	Neurotransmission
β_1	Heart	Increase rate, contractility, conduction
	Adipose tissue	Lipolysis
β_2	Blood vessels	Dilation
	Bronchioles	Dilation
	Kidney	Renin secretion
	Liver	Gluconeogenesis, glycogenolysis
	Endocrine pancreas	Insulin secretion
	Uterus	Relaxation
D_1	Blood vessels	Dilation
D_2	Presynaptic SNS nerve endings	Inhibition of NE release



Parasympathetic Nervous System

M ₁	Skeletal prejunctional nerve endings	Facilitate ACh release
M ₂	Lung–presynaptic PNS nerve endings	Inhibits ACh release
M ₃	Visceral organs Lung smooth muscle, postsynaptic	Increase Bronchoconstriction
N ₁	PNS and SNS ganglion	Ganglionic blockade
N ₂	Skeletal muscle	Muscle contraction



ORGAN SYSTEM	EFFECTOR	SNS STIMULATION	PNS STIMULATION
Cardiovascular	Sinoatrial node	Increase heart rate	Decrease heart rate
	Atrioventricular node	Increase conduction velocity	Decrease conduction velocity
	His-Purkinje fiber	Increase automaticity, conduction	None or minimal
	Myocardial tissue	Increase contractility	None or minimal
	Coronary arterioles	Constriction (α_2), Dilation (β_2)	Dilation and constriction
Gastrointestinal	Salivary glands	Little increase in secretion	Stimulates secretion
	Gall bladder	Relaxation	Contraction
	Smooth muscle	Inhibits	Stimulates
	Sphincters	Constricts	Relaxes
Ophthalmologic	Iris	Mydriasis	Miosis
	Ciliary muscle	Relaxation (for far vision)	Contraction (for near vision)
Pulmonary	Bronchial smooth muscle	Relaxation	Contraction
Genitourinary	Bladder walls	Relaxes smooth muscle	Smooth muscle contraction
	Bladder sphincter	Contraction	Relaxation
	Ductus deferens, seminal vesicle, prostatic and uterine musculature	Contraction of smooth muscle	Vasodilation and erection
Thermal regulation	Blood vessels	Constriction	Dilation
	Sweat glands	Diaphoresis (postganglionic sympathetic fibers are cholinergic)	Little effect

Autonomic reflexes

- Thermogenesis reflex
- Baroreceptor reflex
- Chemoreceptor reflex
- Bainbridge reflex
- Bezold –Zarisch reflex
- Oculocardiac reflex
- Valsava manœuvre
- Cushing reaction

- The ANS receives input from parts of the CNS that process and integrate stimuli from the body and external environment.
- **Through direct projection to preganglionic neurons-**
 - Paraventricular nuclei of hypothalamus,
 - Nucleus of the solitary tract,
 - reticular formation,
 - Parabrachial nuclei
 - Ventrolateral Medulla,
 - Medullary Raphe
 - amygdala, hippocampus, and olfactory cortex.

- **Through indirect Projections:**

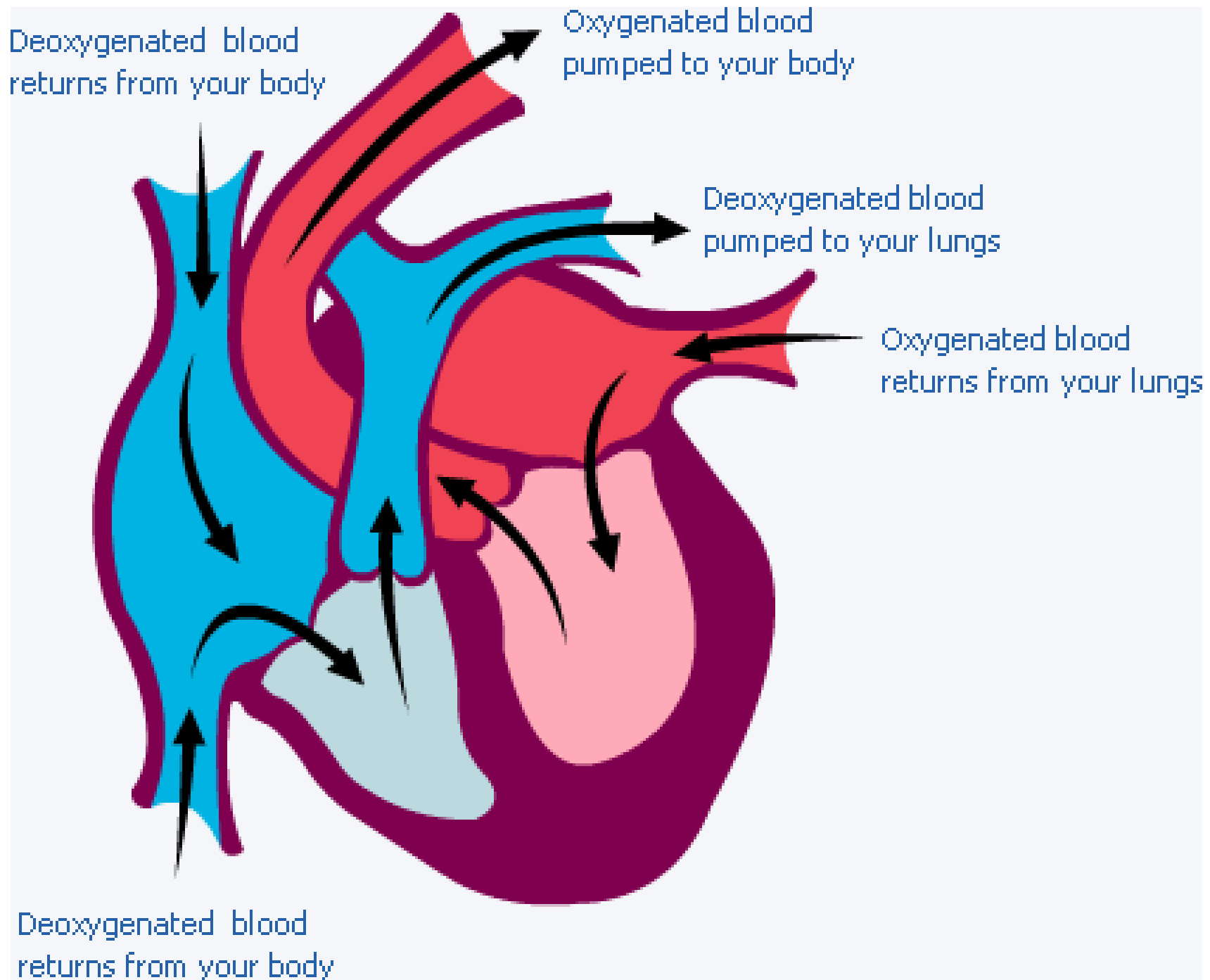
- Cerebral cortex
- Amygdala
- Periaqueductal grey Matter

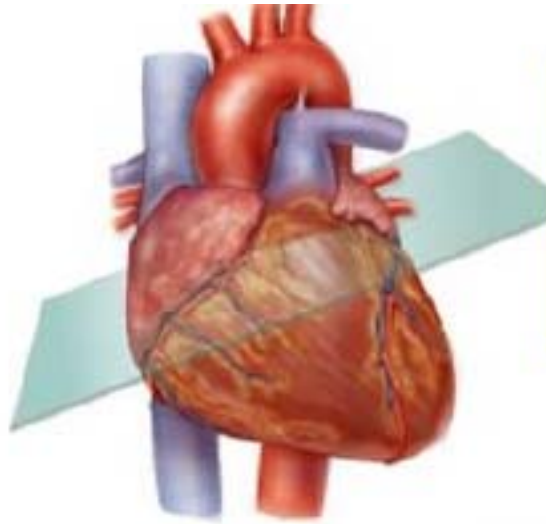
CARDIOVASCULAR SYSTEM

“It has been shown by reasons and experiments that blood by the beat of the ventricles flows through the lungs and heart and is pumped to the whole body... the blood in the animal body moves around in the circle continuously, and ..the action of the heart is to accomplish this by pumping. This is the only reason for the motion and beat of the heart.”

-William Harvey (1628)

- Two pumps – Operating in series
- All four valves lie in a plane within a connective tissue skeleton which separates atria and ventricle
- Mitral, tricuspid and aortic valves surround a fibrous triangle “central fibrous tendon/body”





The **fibrous skeleton** of the heart is located between the atria and the ventricles, and is formed from dense irregular connective tissue. It is electrically non-conductive!

Posterior

Right
atrioventricular
valve

Aortic semilunar
valve

Openings to coronary
arteries

Pulmonary semilunar valve

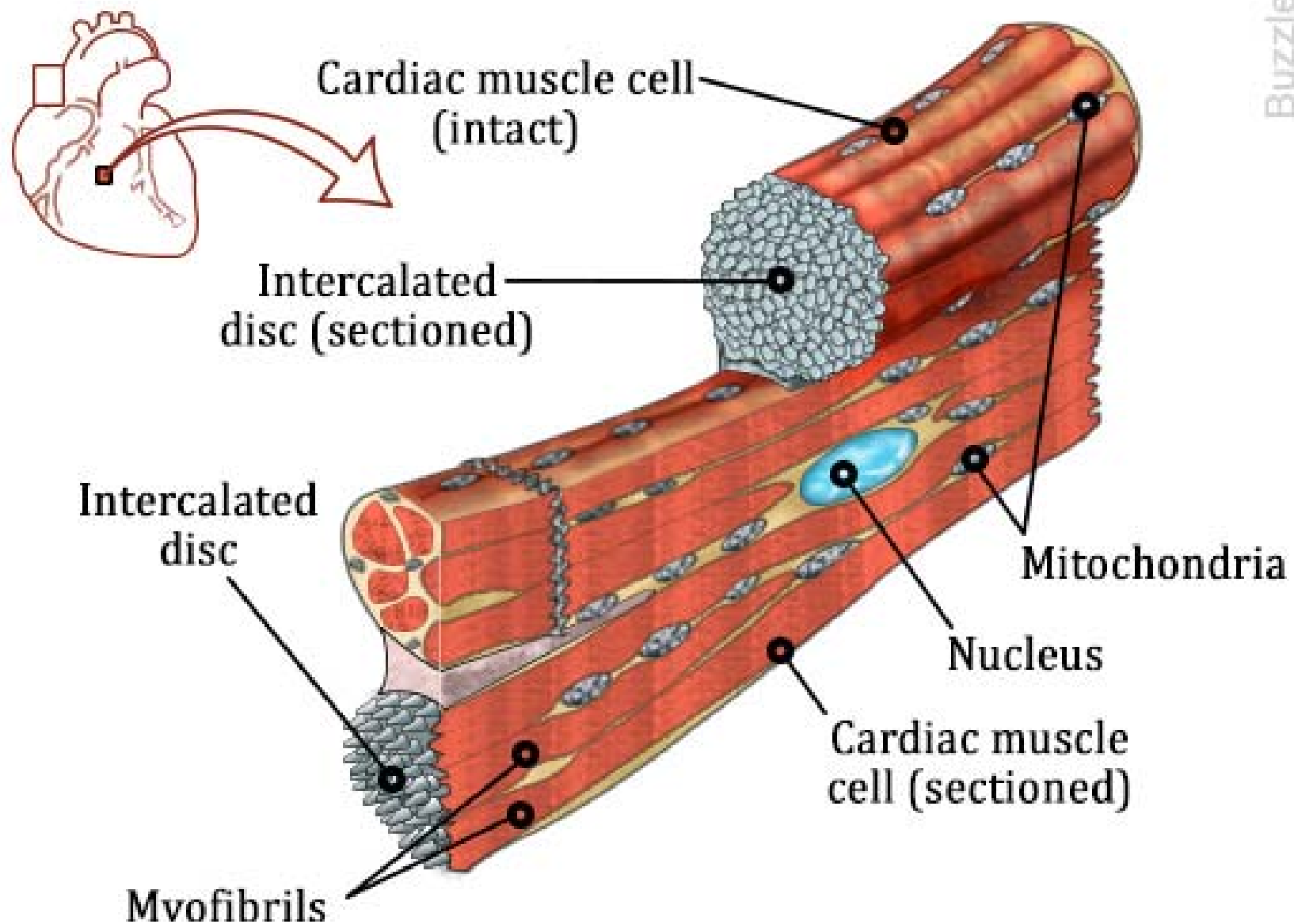
Left
atrioventricular
valve

Fibrous
skeleton

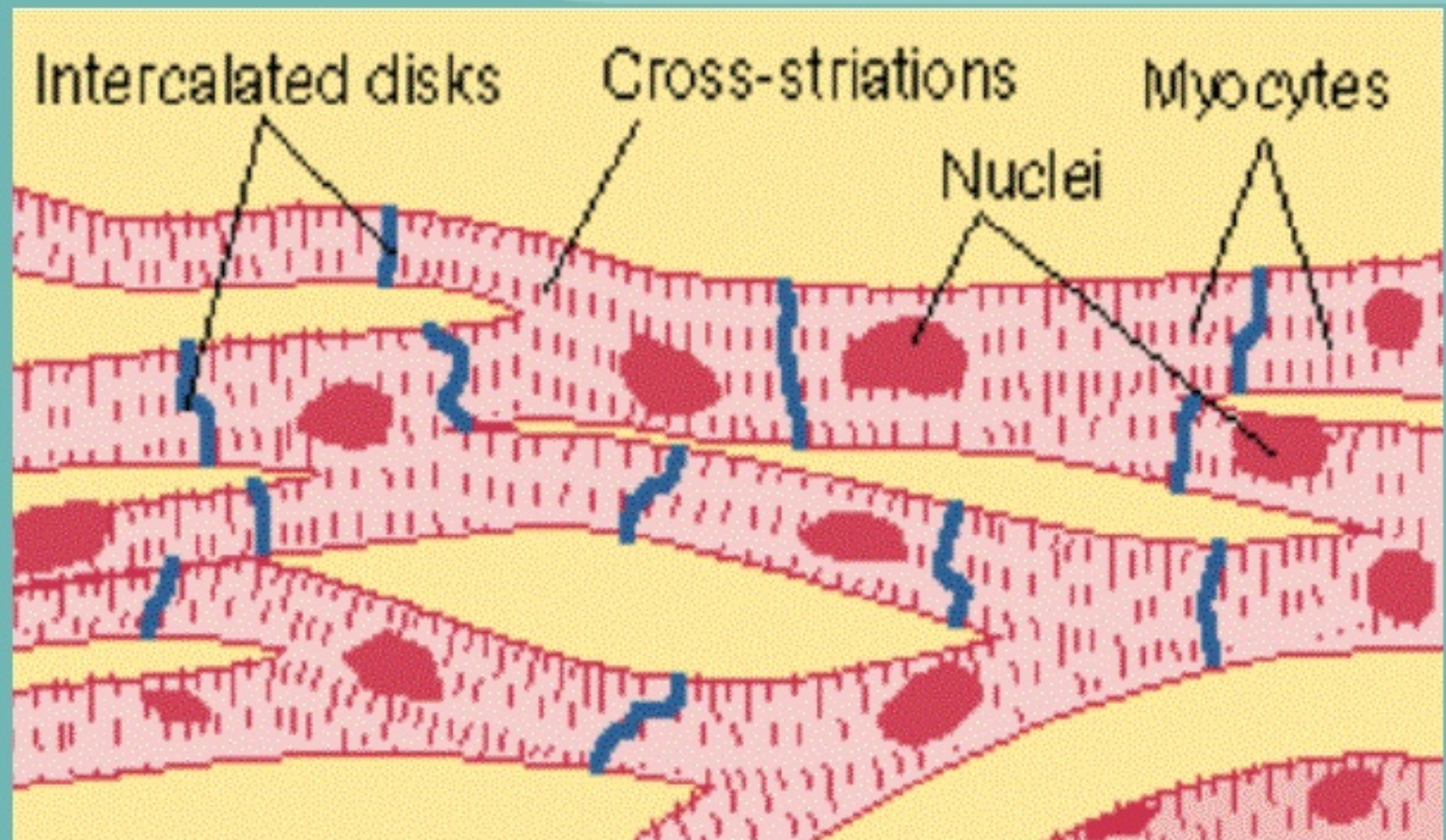


Anterior

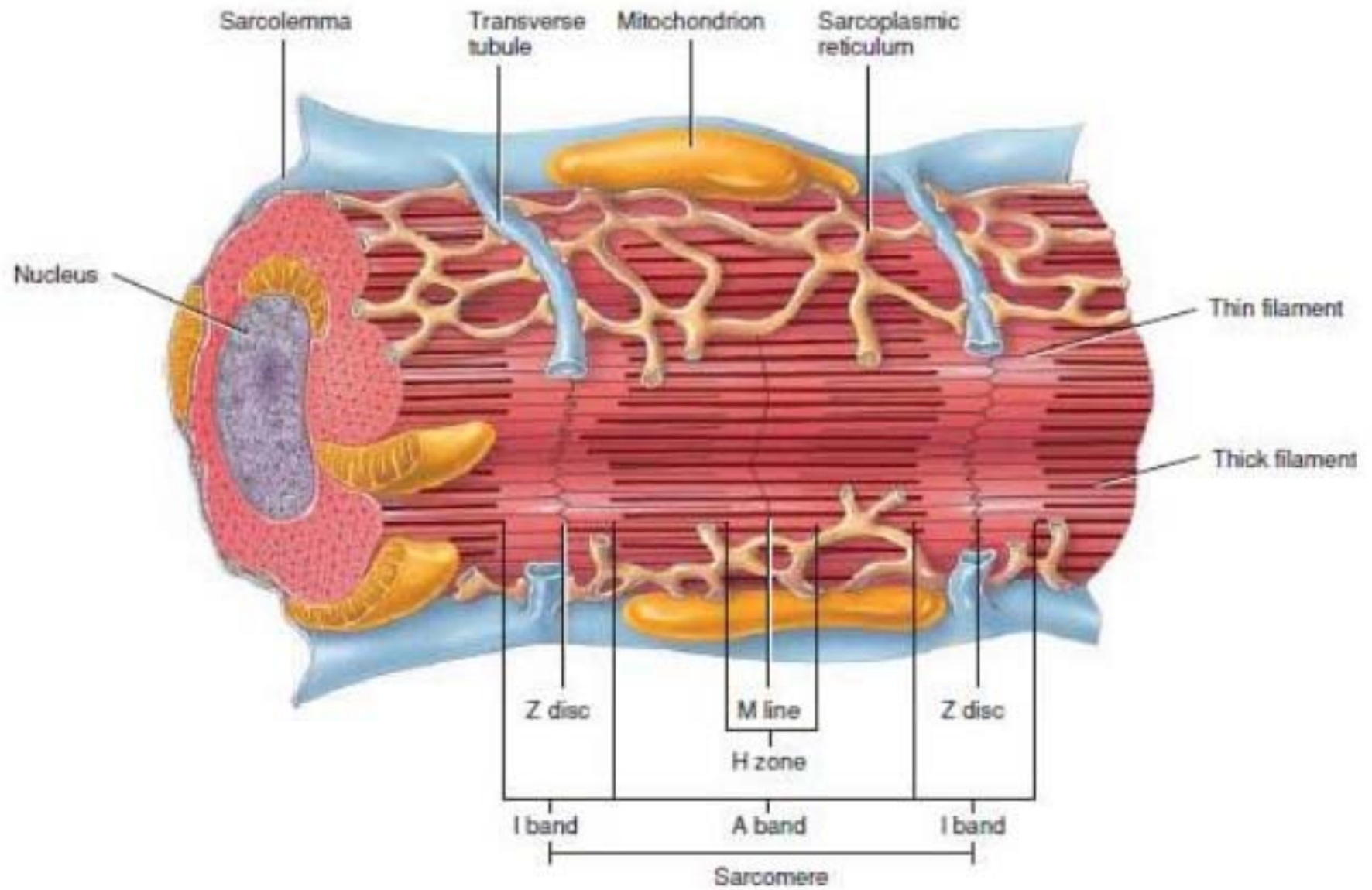
Cardiac Muscle Cross Section



Cardiac Muscle



(a) Cardiac muscle fibers

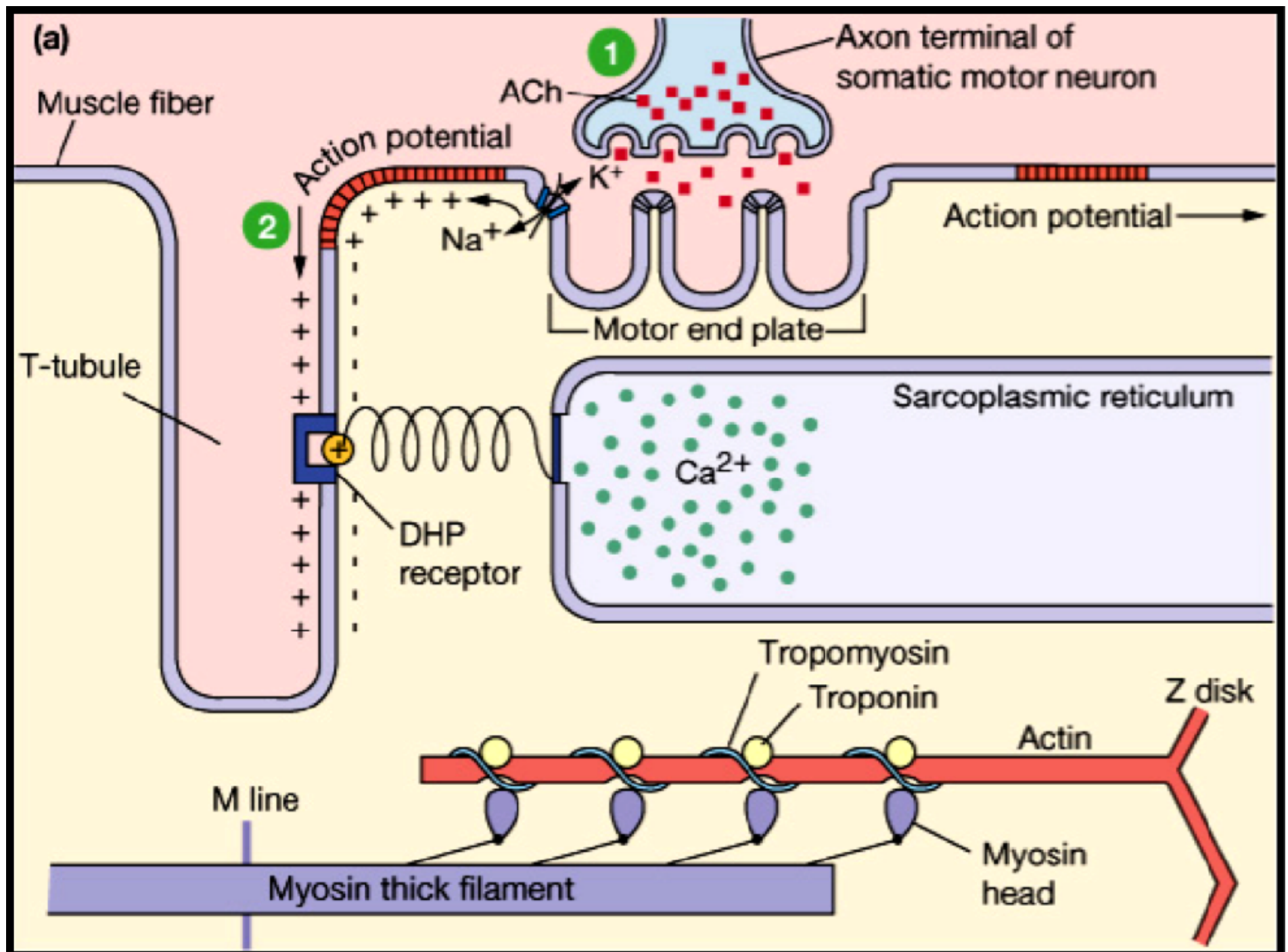


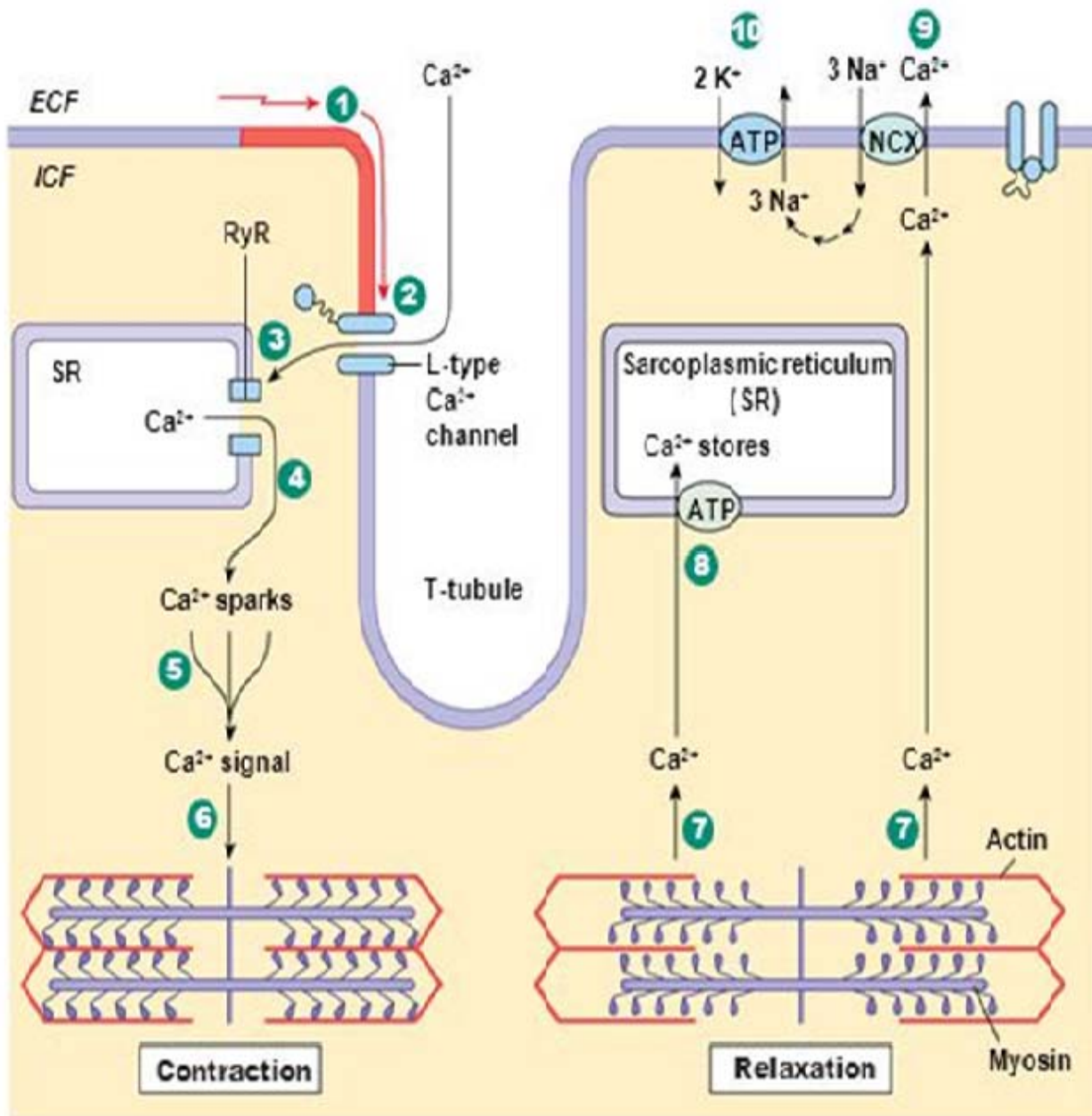
CARDIAC MUSCLE

- **Epicardium**- A layer of squamous cells overlying a network of fibroelastic connective tissue .
- Continuous with inner layer of pericardium.
- **Endocardium** – A squamous cell layer, a mesh of collagen and elastic fibres and rudimentary smooth muscle.
- **Myocardium** – Cardiac myocytes,
 - 70% are non myocyte small cells like vascular smooth muscle, endothelial cells and fibroblasts.

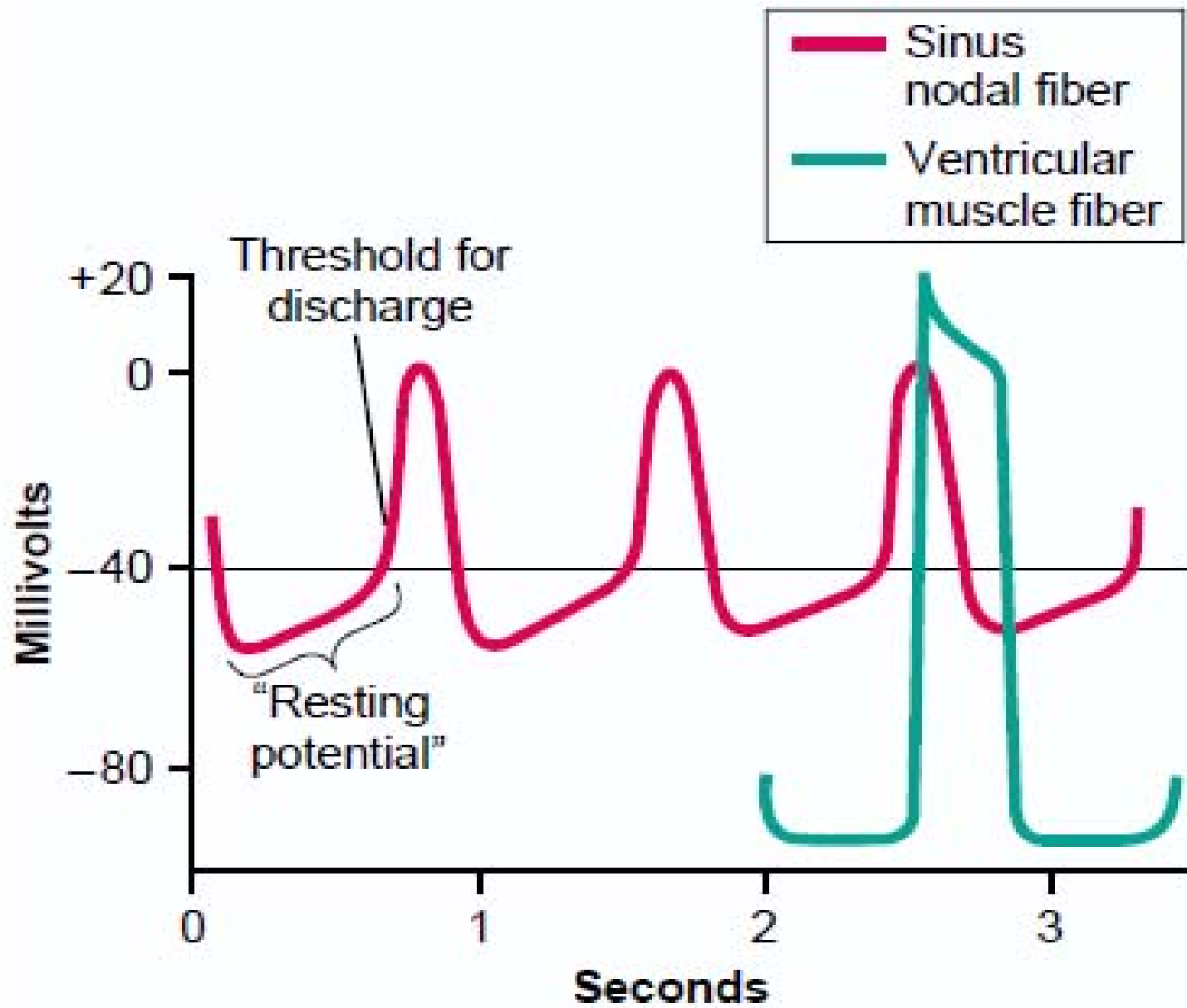
- **Fibroblast** – Secrete and maintain connective tissue → contribute to heart's tensile strength and stiffness.
- **Cardiac Myocytes**
 - **Working Myocytes**- Specialized contractile cardiac muscle
 - **Purkinje Fibres** – large pale, glycogen rich
 - Specialized for rapid conduction.
 - **Nodal cells** – Contain fewer myofilaments
 - **Round cells** – Probably the pacemakers
 - **Slender / Transitional cells** – conduct the impulse within the node and nodal margin

- **Intercalated disc** – Mechanical and electrical connection between cardiac muscle
- **Mechanical** _ fascia adherens
 - Sarcomeric actin is connected to cytoskeletal actin and desmosomes connecting intermediate filament
- **Gap junction** – least resistance for free diffusion of ions and small molecules
- **Cardiac myocytes** – depend on oxidative metabolism for energy
- **Conducting tissues** – have developed anaerobic energy metabolism.
- Pace maker cells
- Overdrive suppression – due to overactivity of Na-K pump

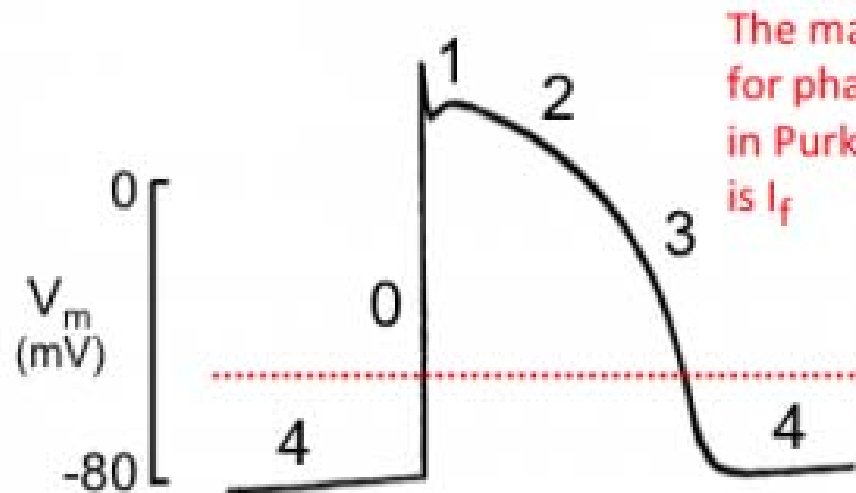




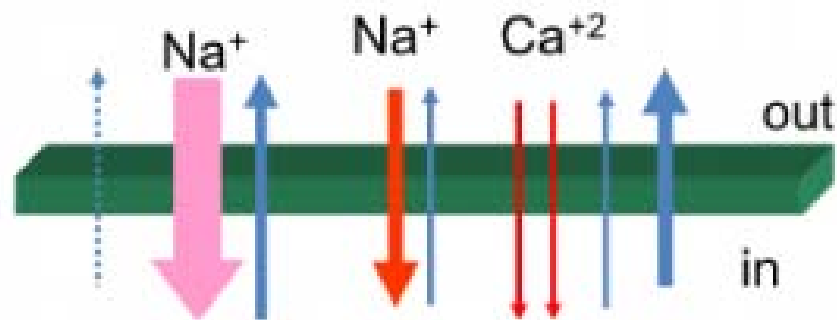
- 1 Action potential enters from adjacent cell.
- 2 Voltage-gated Ca^{2+} channels open. Ca^{2+} enters cell.
- 3 Ca^{2+} induces Ca^{2+} release through ryanodine receptor-channels (RyR).
- 4 Local release causes Ca^{2+} spark.
- 5 Summed Ca^{2+} sparks create a Ca^{2+} signal.
- 6 Ca^{2+} ions bind to troponin to initiate contraction.
- 7 Relaxation occurs when Ca^{2+} unbinds from troponin.
- 8 Ca^{2+} is pumped back into the sarcoplasmic reticulum for storage.
- 9 Ca^{2+} is exchanged with Na^+ by the NCX antiporter.
- 10 Na^+ gradient is maintained by the $\text{Na}^+-\text{K}^+-\text{ATPase}$.



Purkinje Fiber

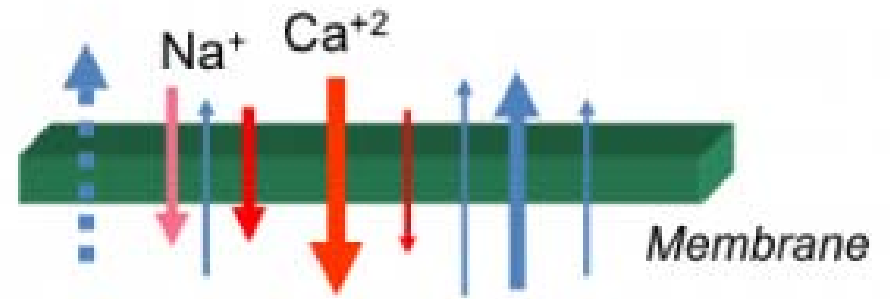
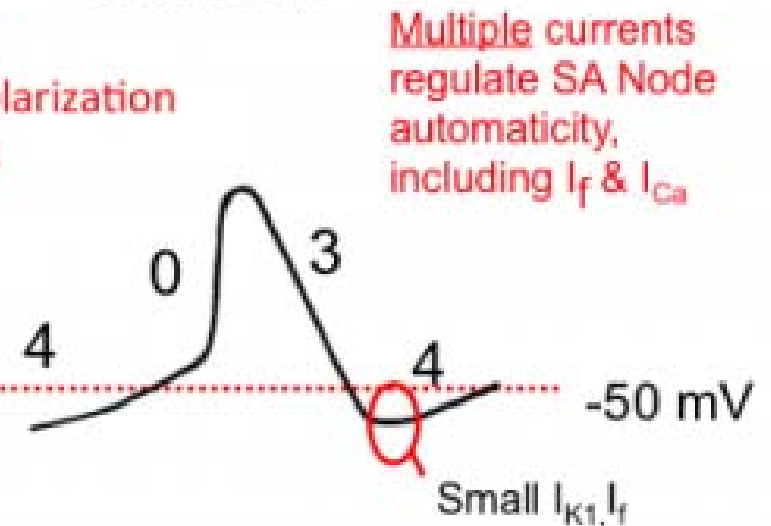


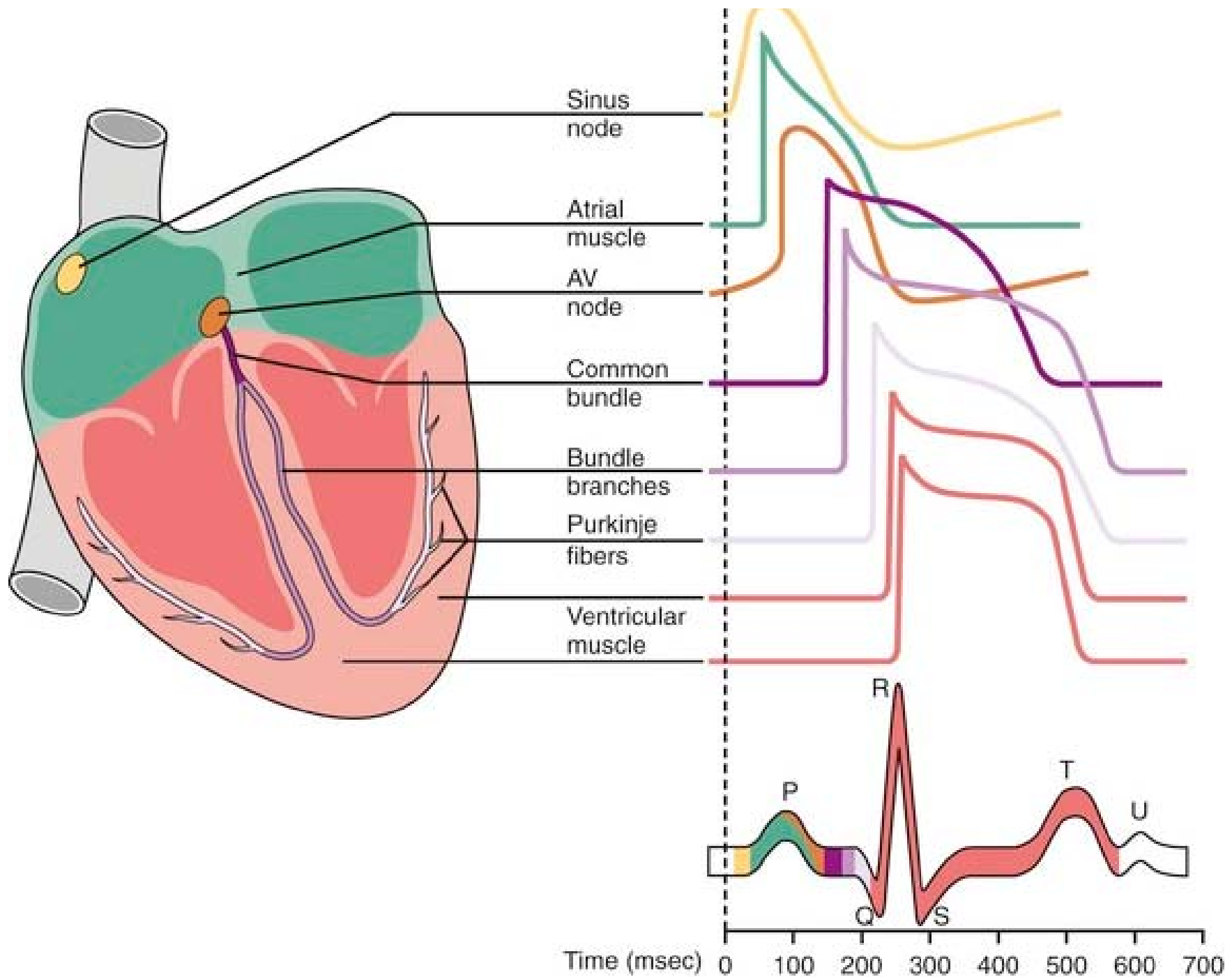
The main source for phase 4 depolarization in Purkinje fibers is I_f



Currents that regulate pacemaking

SA Node





CHANNEL	CHARACTERISTICS
<i>Sodium Channels</i>	
Fast Na^+	Phase 0 depolarization of non-pacemaker cardiac action potentials
Slow Na^+	"Funny" pacemaker current (I_f) in cardiac nodal tissue
<i>Potassium Channels</i>	
Inward rectifier (I_{ir} or I_{K1})	Maintains phase 4 negative potential in cardiac cells and contributes to late repolarization
Transient outward (I_{to})	Contributes to phase 1 of non-pacemaker cardiac action potentials
Delayed rectifier (I_{Kr})	Phase 3 repolarization of cardiac action potentials
ATP-sensitive ($I_{K, ATP}$)	K_{ATP} channels; inhibited by ATP; therefore, open when ATP decreases during hypoxia; in vascular smooth muscle, adenosine removes the ATP inhibition and opens these channels, producing hyperpolarization and vasodilation.

Acetylcholine-
activated ($I_{K,ACh}$)

Activated by acetylcholine; Gi-protein coupled

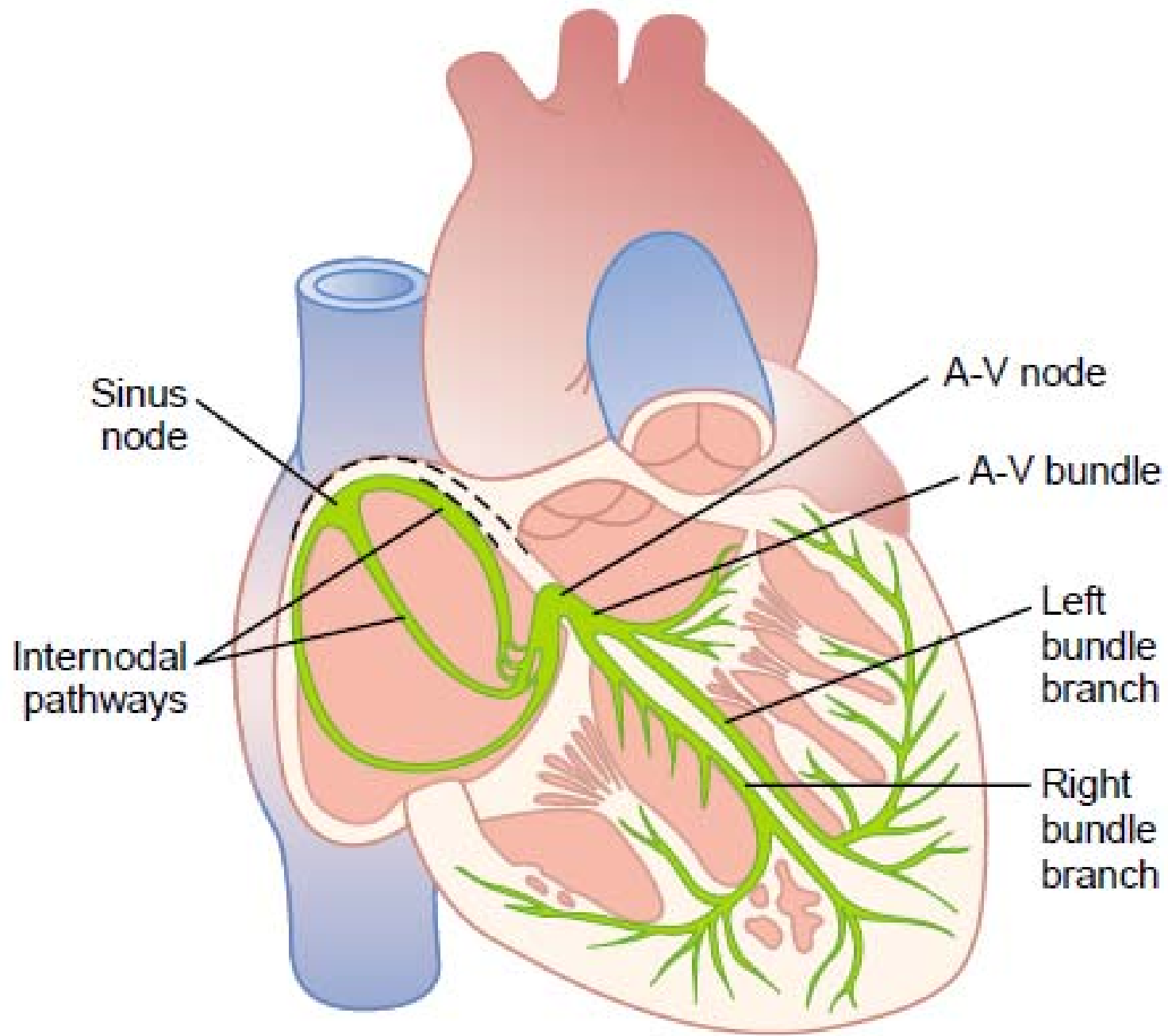
Calcium-

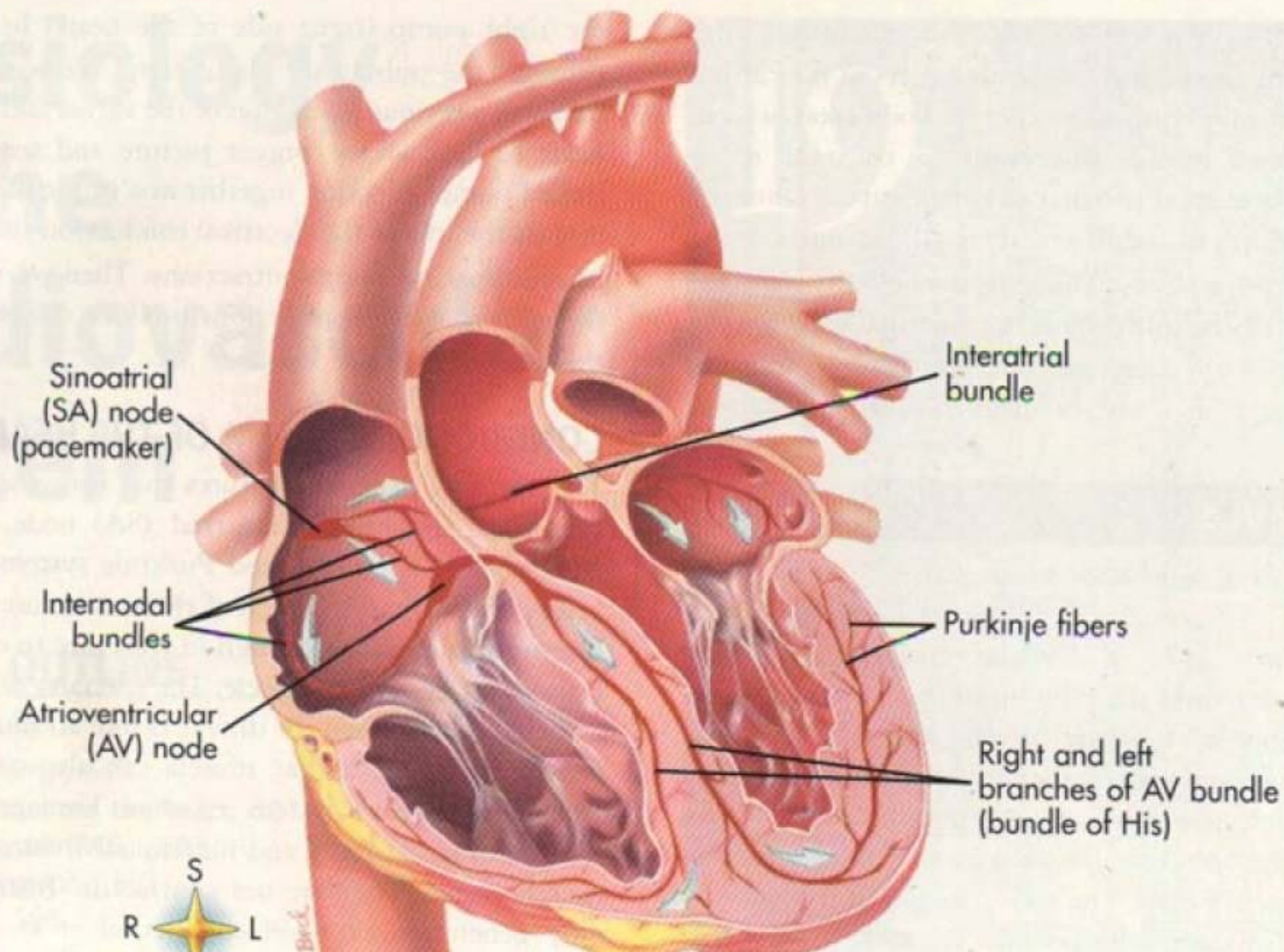
activated ($I_{K,Ca}$ or BK_{Ca}) Open in response to Ca^{++} influx in vascular smooth muscle

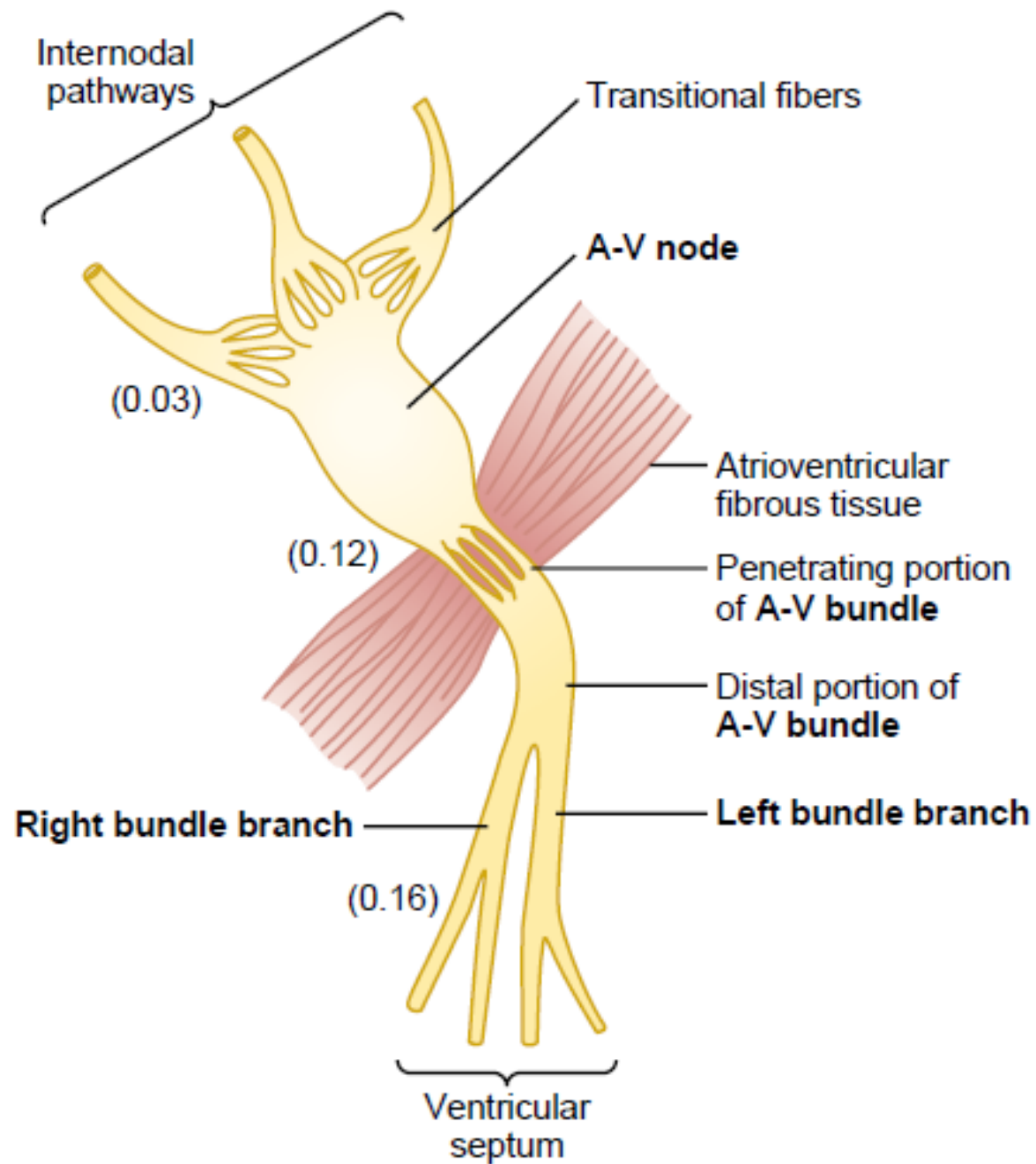
*Calcium
Channels*

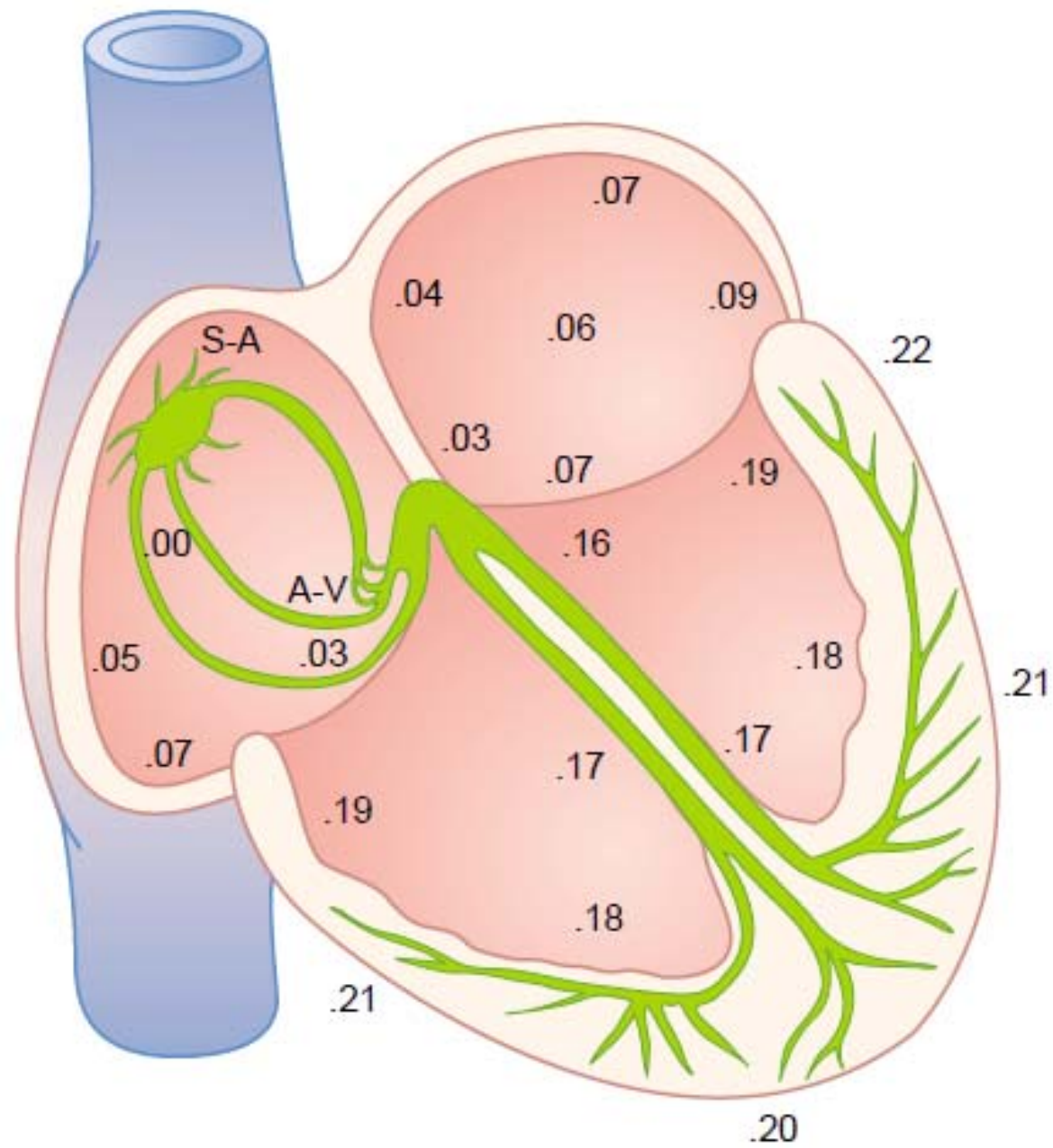
L-type (I_{Ca-L}) Slow inward, long-lasting current; phase 2 non-pacemaker cardiac action potentials and late phase 4 and phase 0 of SA and AV nodal cells; important in vascular smooth muscle contraction

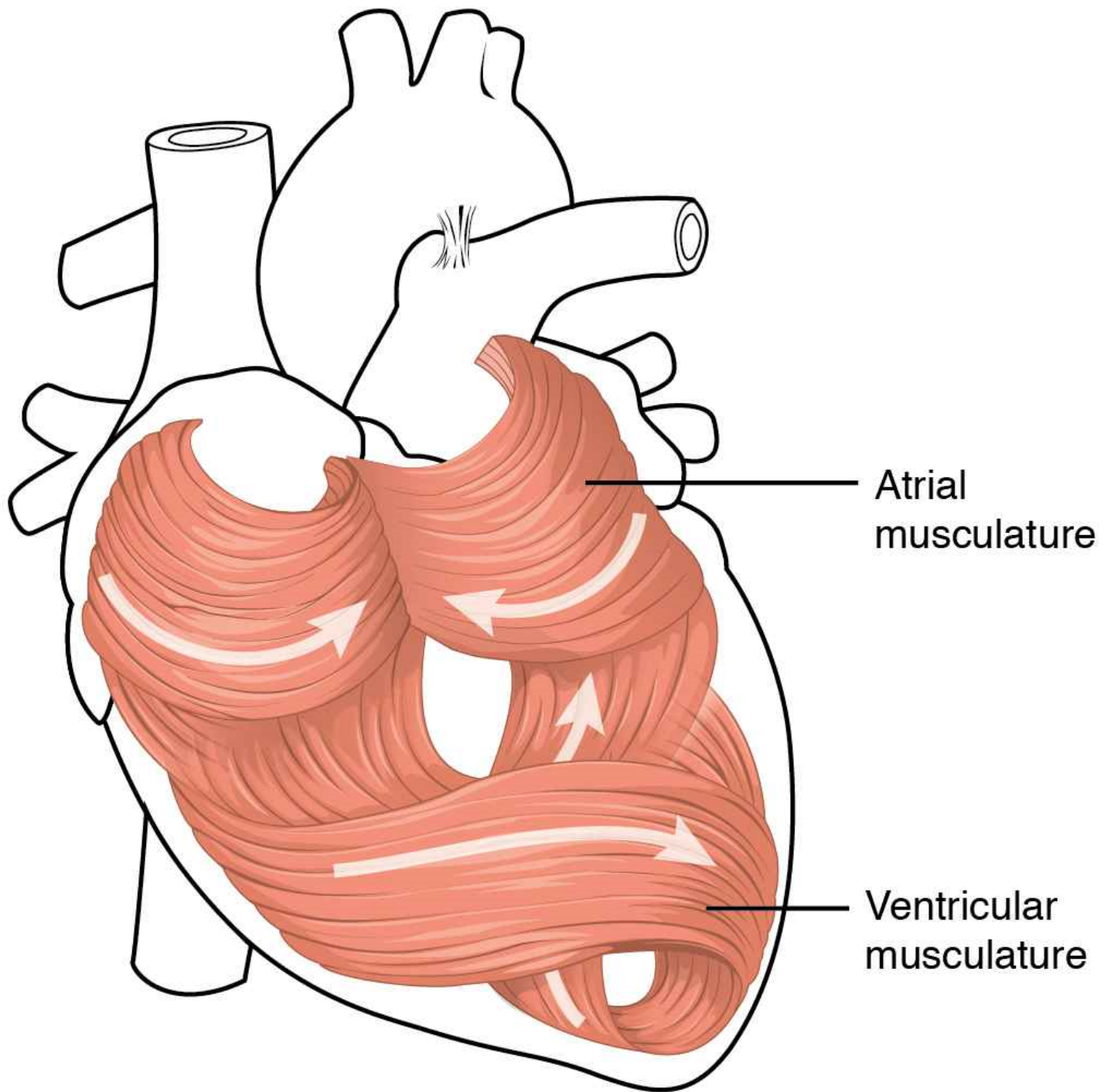
T-type (I_{Ca-T}) Transient current that contributes to early phase 4 pacemaker currents in SA and AV nodal cells











CONDUCTING SYSTEM

Tissue	Conduction Velocity	Time of Impulse Delay
S A Node		0
Atrial Muscle	0.3m/sec	.03sec
Inter nodal fibres	1m/sec	
AV node	.05m/sec	.09 sec
Penetrating part of AV node/Bundle of His	1m/sec	.04 sec
Purkinje Fibres	1.5– 4 m/sec	.16 sec
Termination of Purkinje Fibres		.03 sec
Ventricular muscle	1m/sec	.03 sec

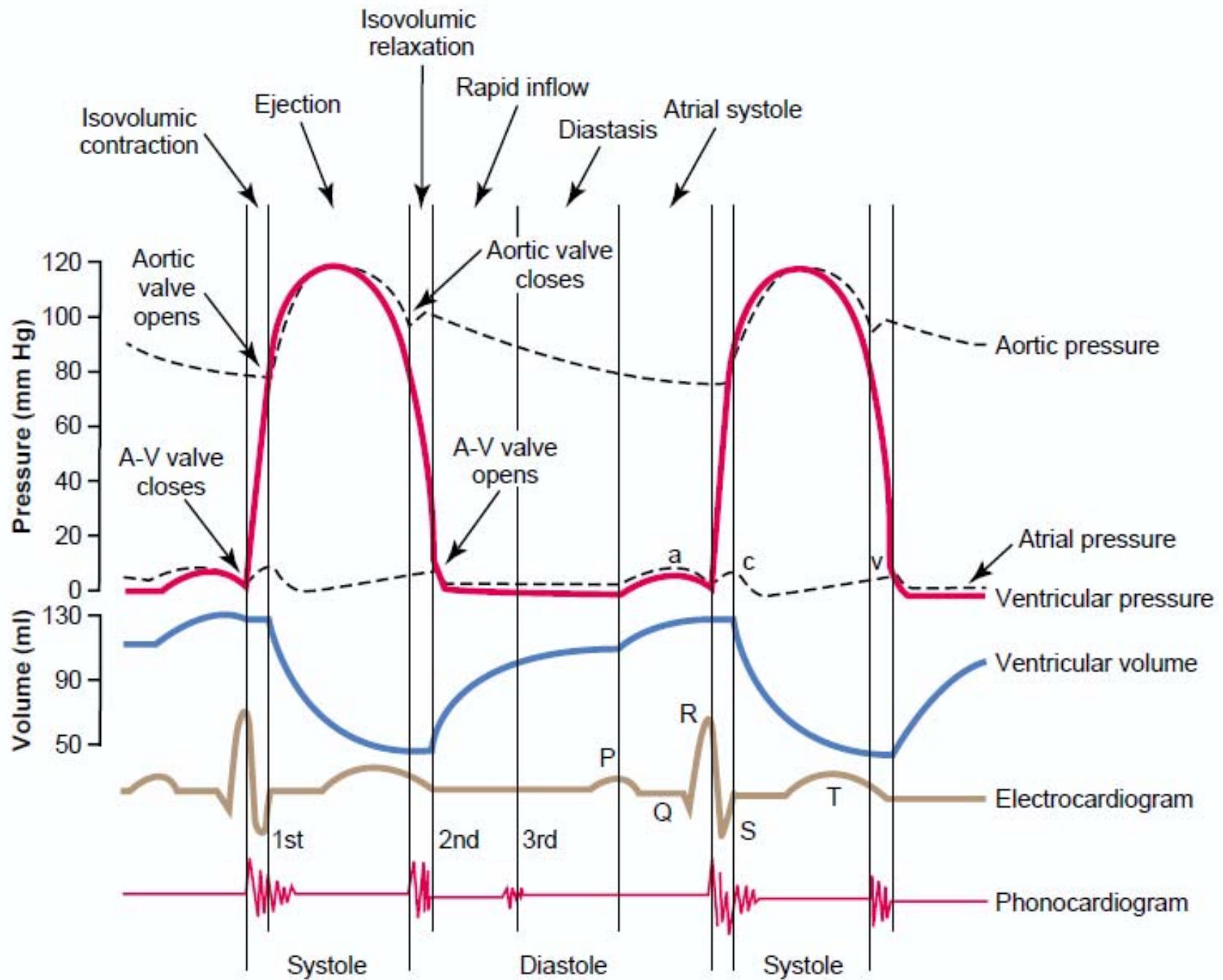
- Right atrium → Left atrium --- Bachman's Bundle
- Internodal Pathway

EFFECT OF ADRENALINE ON HEART

- Effect on
 - L-type Ca^{++} Channels
 - Delayed Rectifier channels
 - Hyperpolarization – activated Cation channels (I_f)
- Also increases intracellular Ca^{++} in atrial, ventricular and AV cells

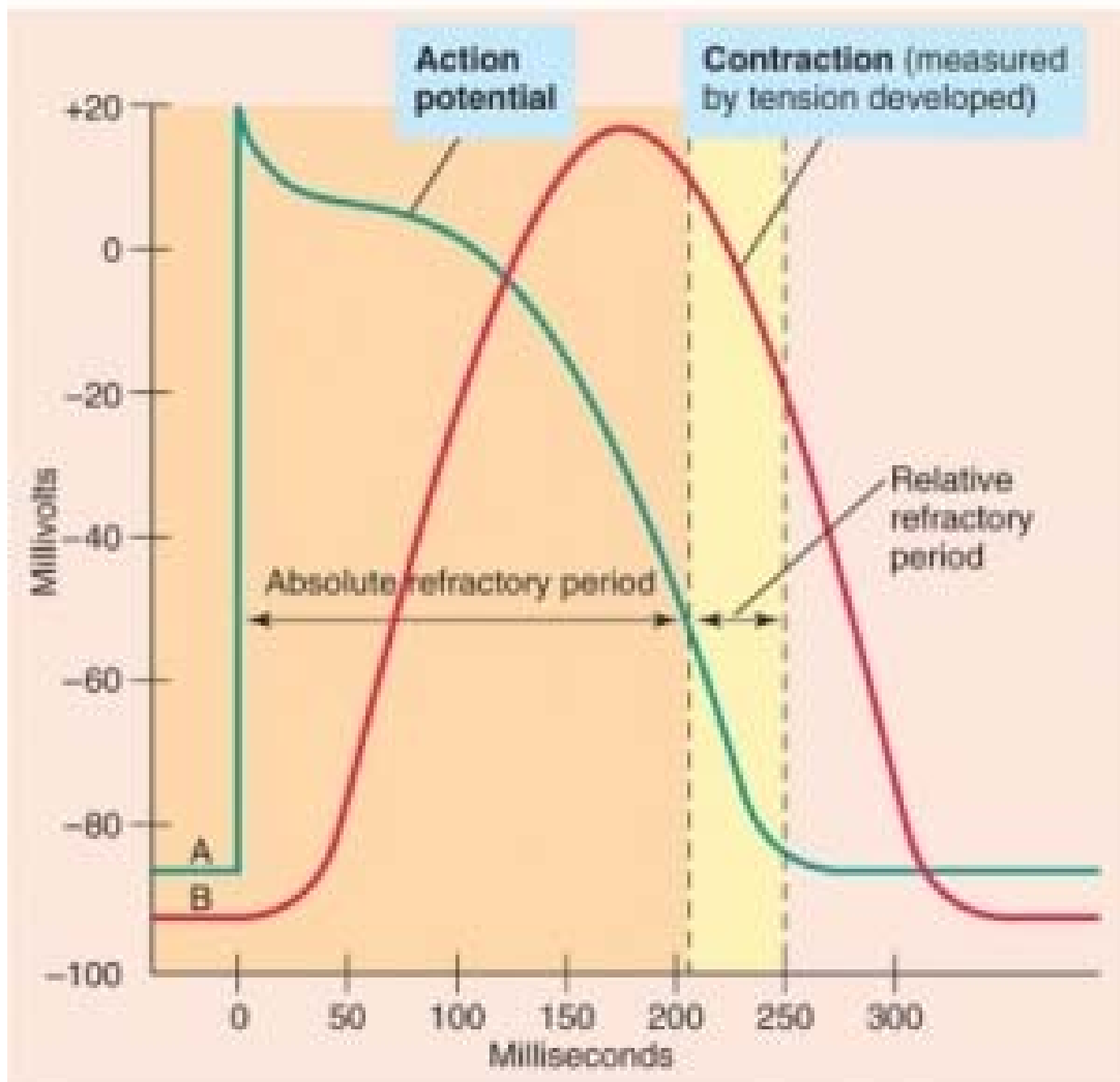
ACETYLECHOLINE

- Increases K⁺ conductance

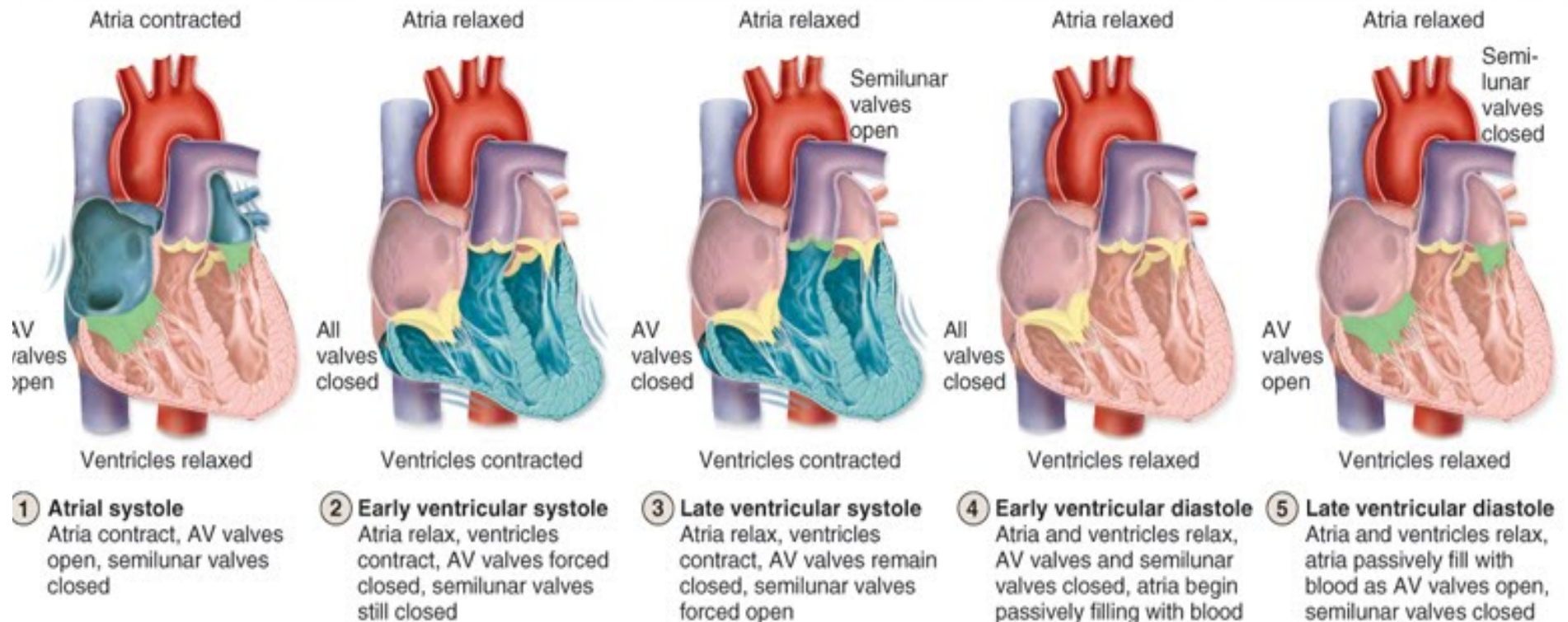


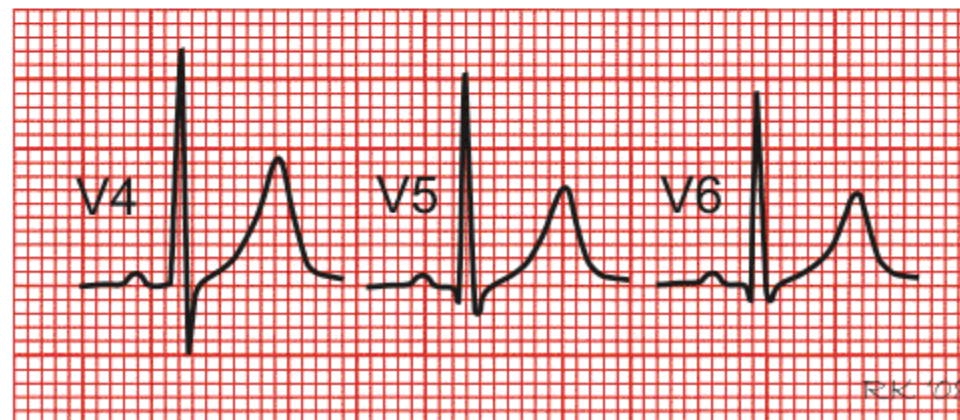
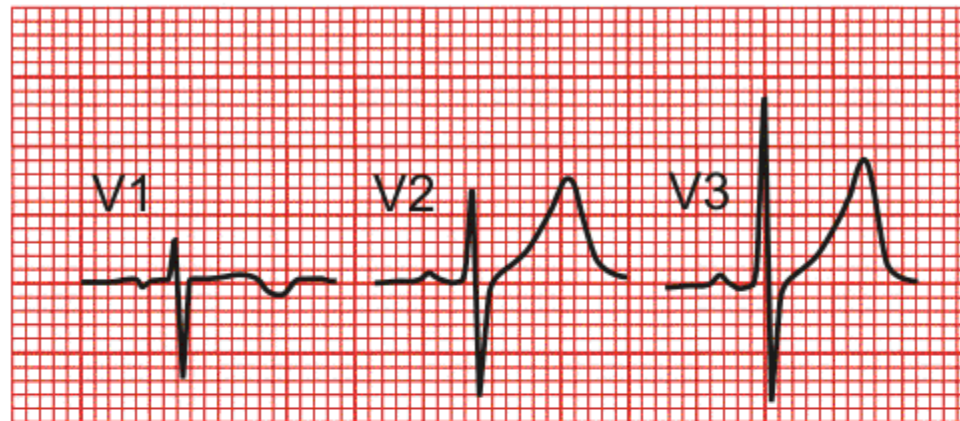
- Isovolumetric contraction- 0.05s
- Rapid ejection – 0.20s
- Protodiastole – 0.05s

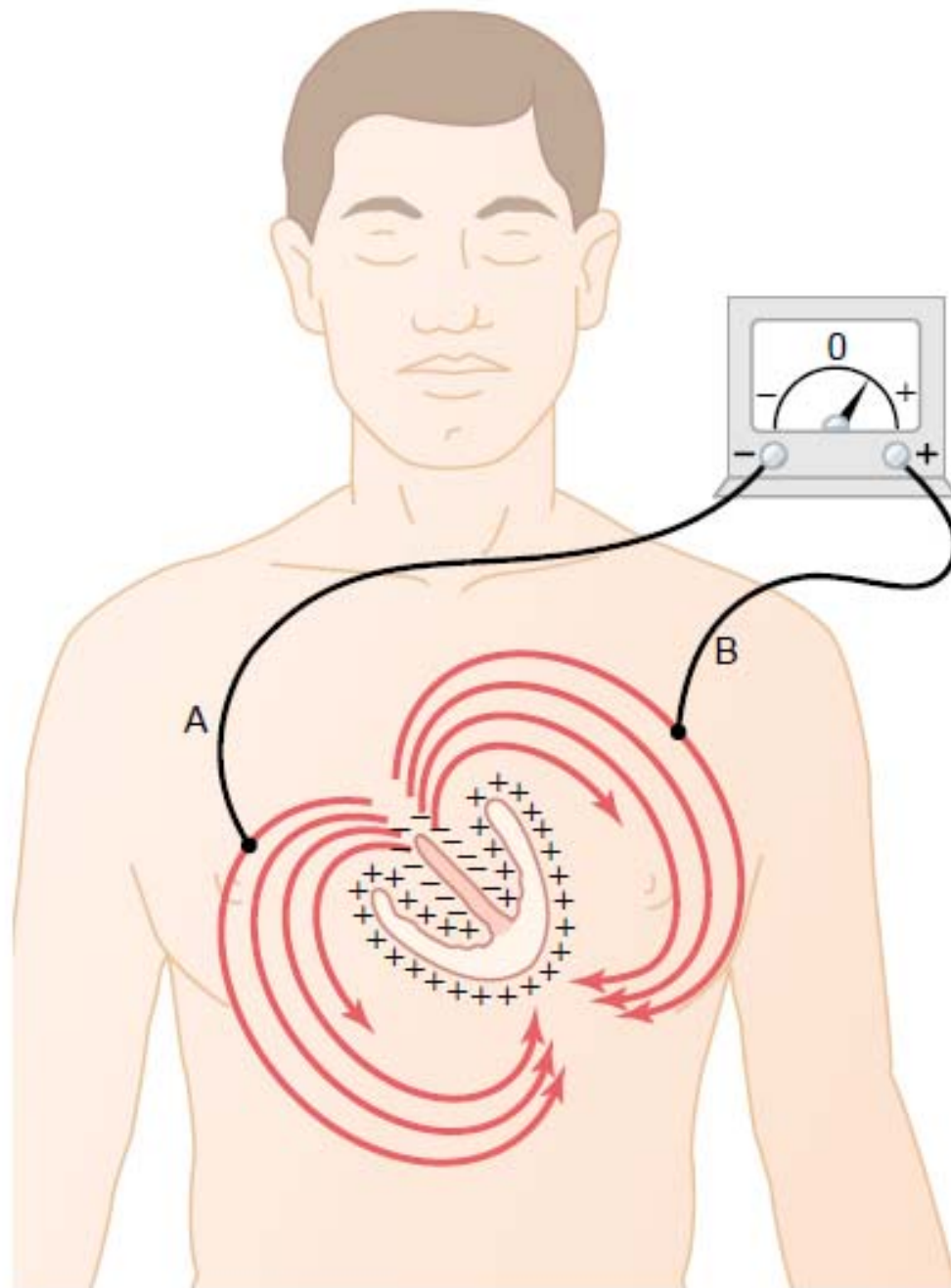
- Isovolumetric relaxation – 0.05s
- Rapid filling – 0.05s
- Diastasis – 0.30s
- Atrial systole – 0.10s

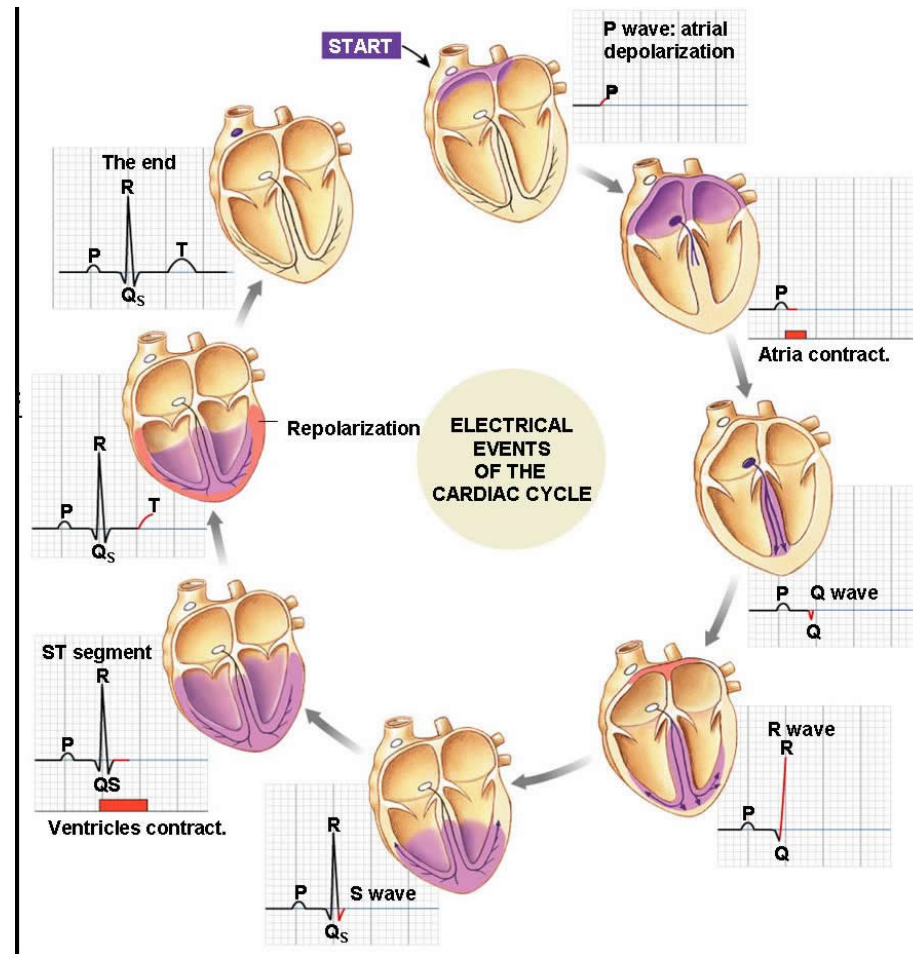


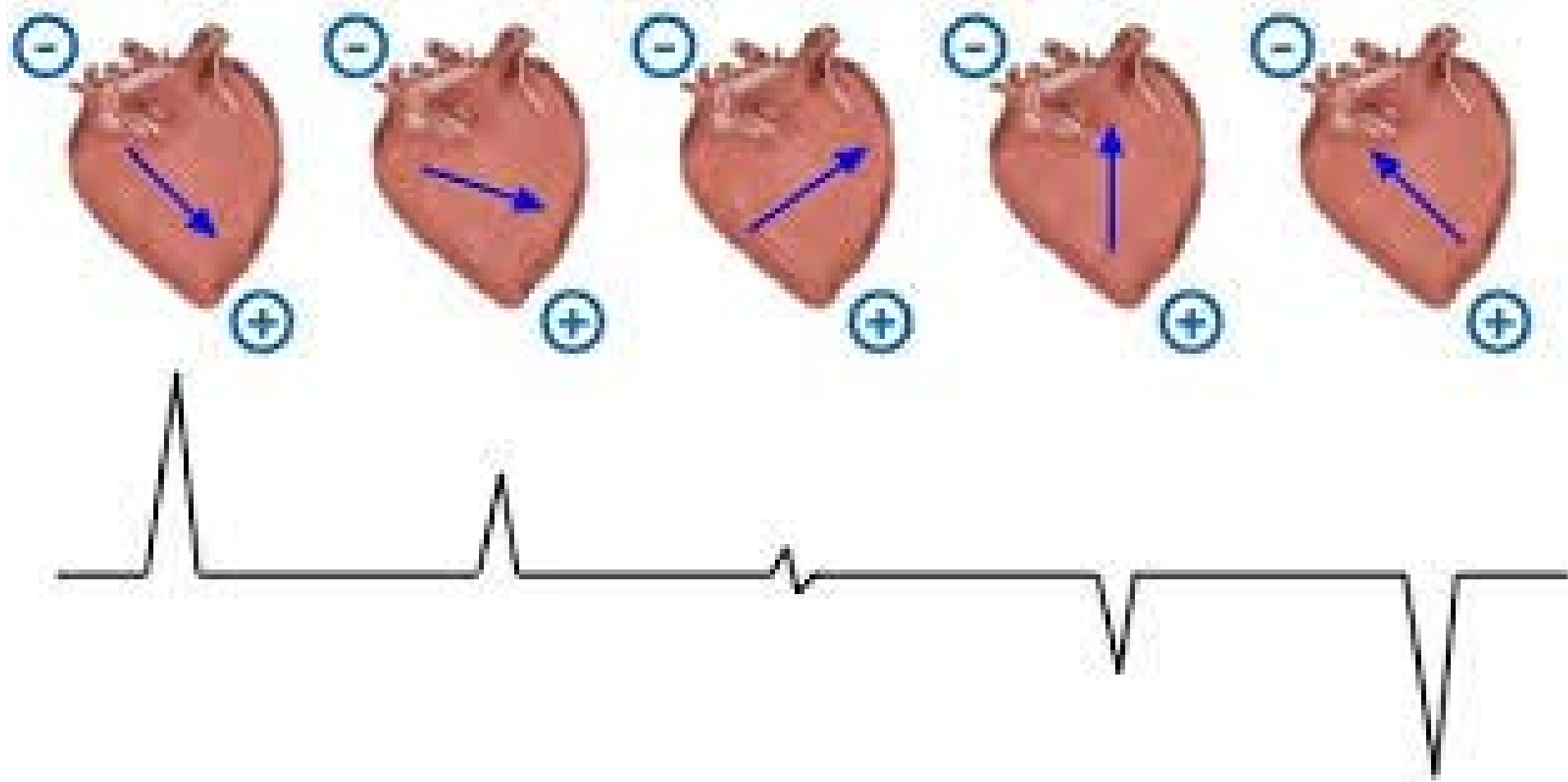
Phase \ Structure	Atrial systole	Early ventricular systole	Late ventricular systole	Early ventricular diastole	Late ventricular diastole
Atria	Contract	Relax		Relax	
Ventricles	Relax	Contract		Relax	
AV valves	Open	Closed		Open	
Semilunar valves	Closed	Open		Closed	



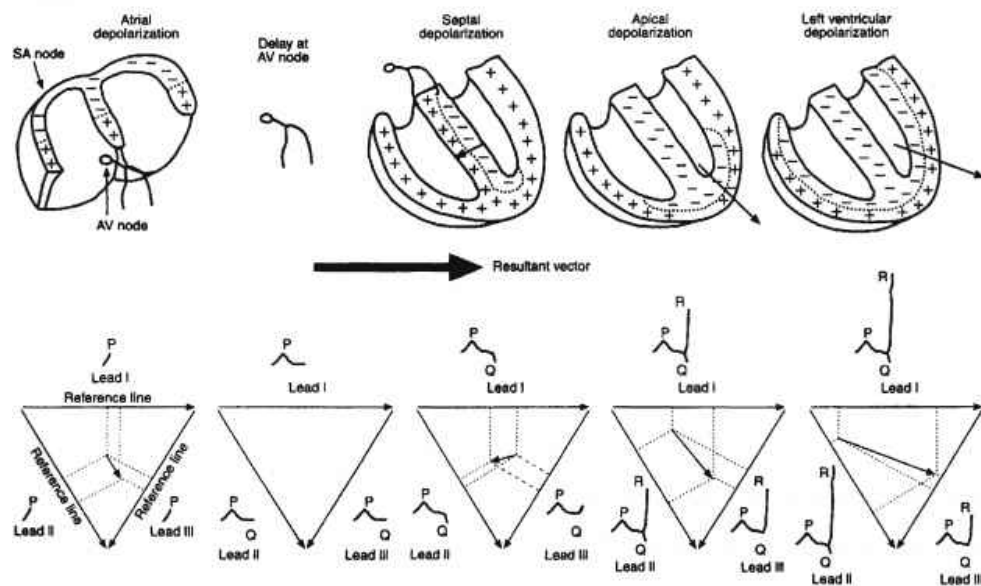




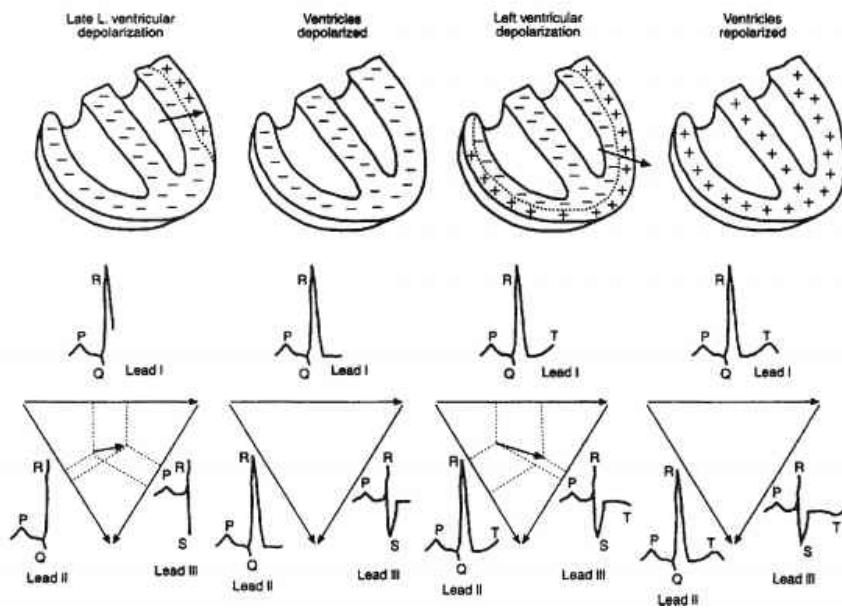


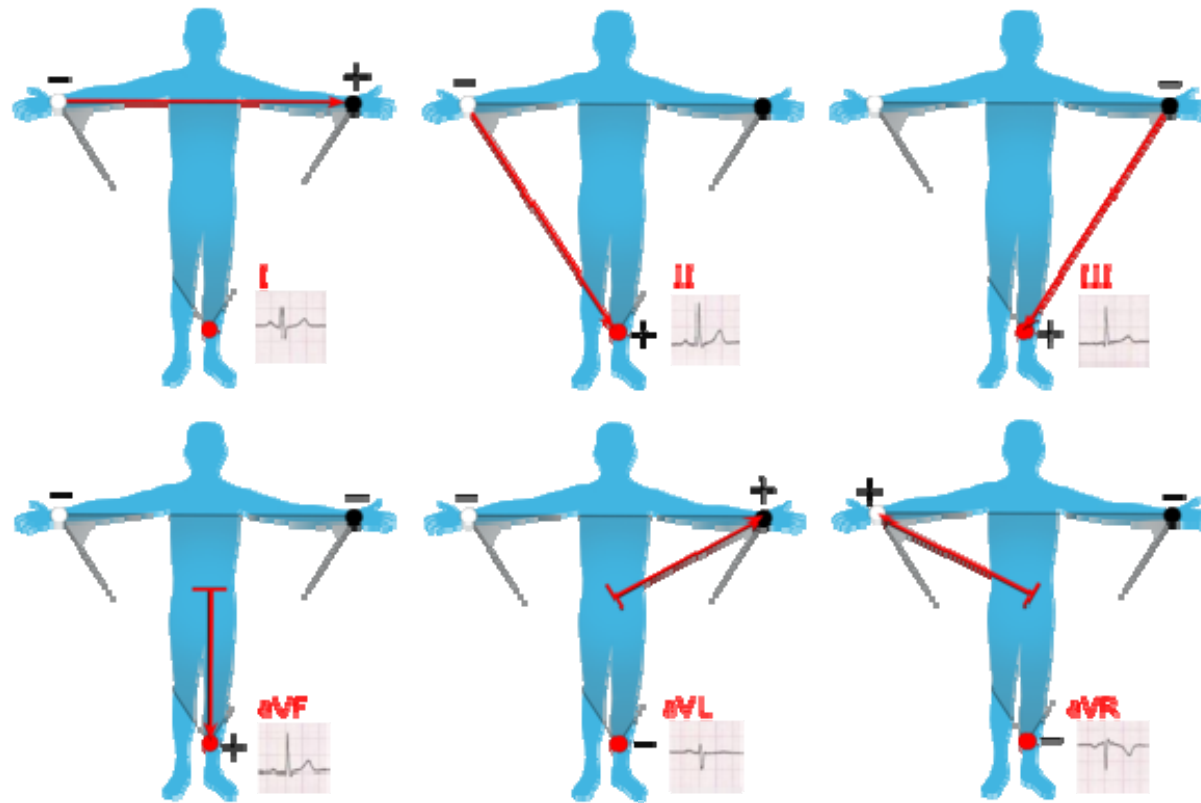


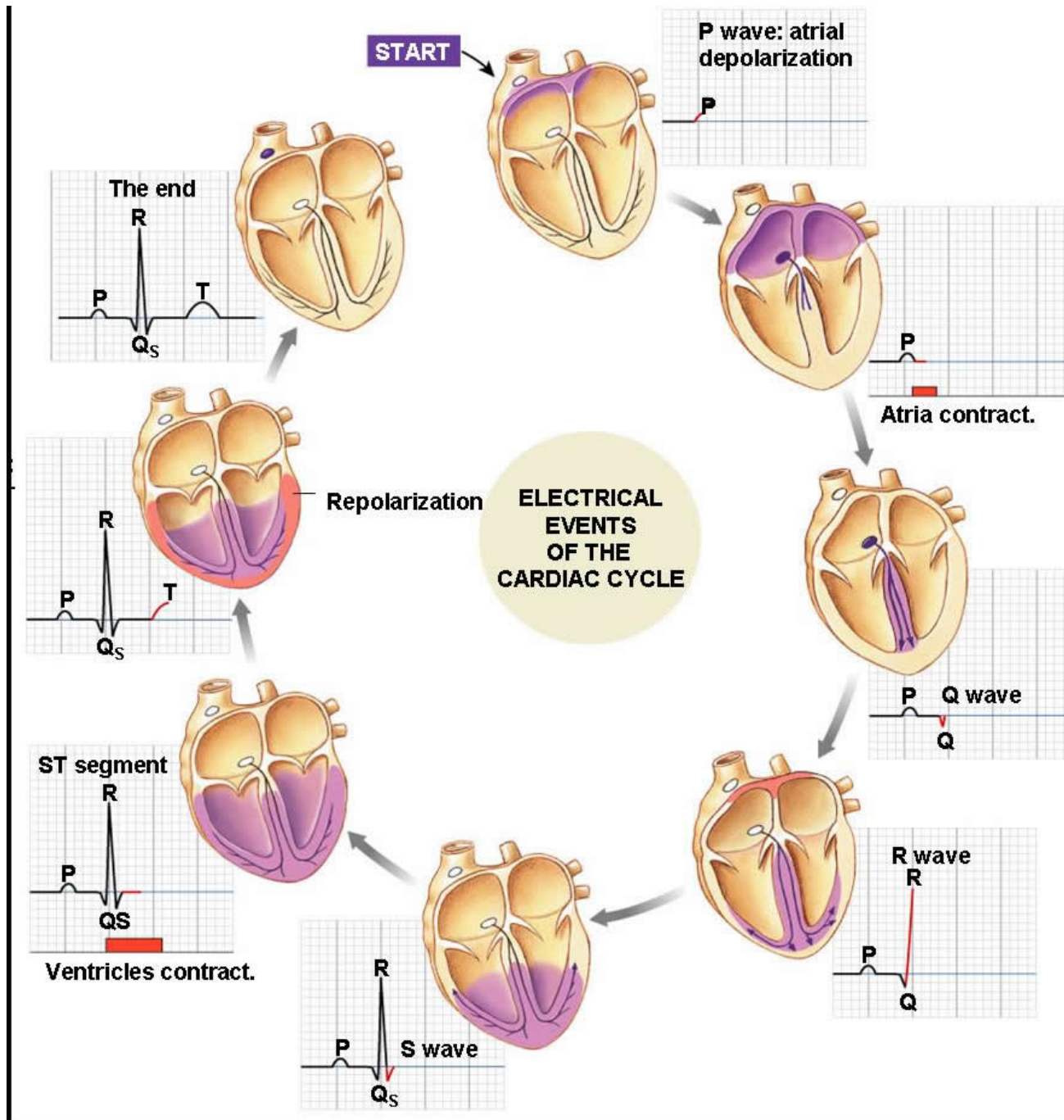
Progression of depolarization



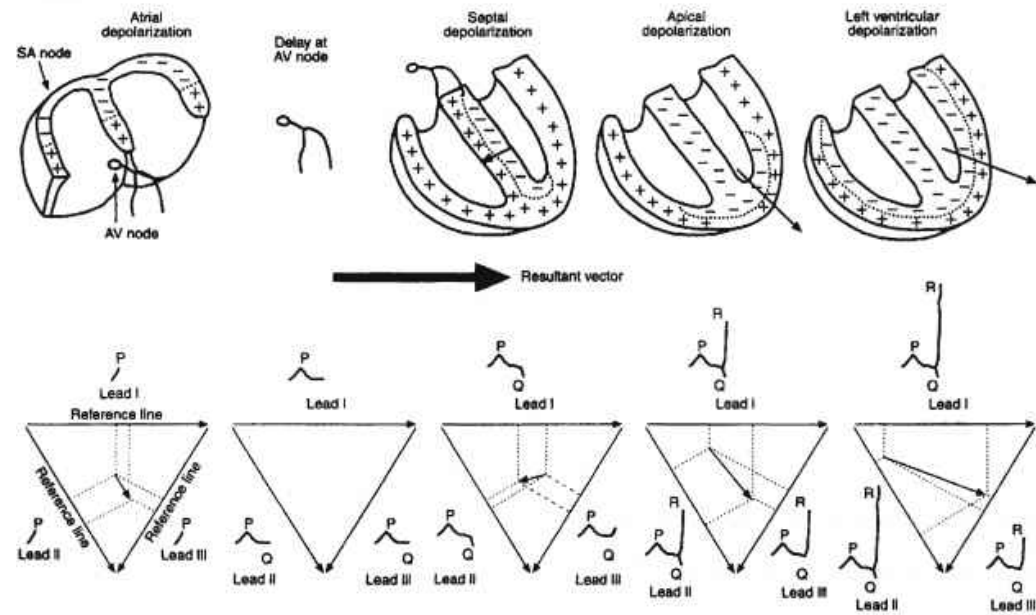
End of depolarization and repolarization



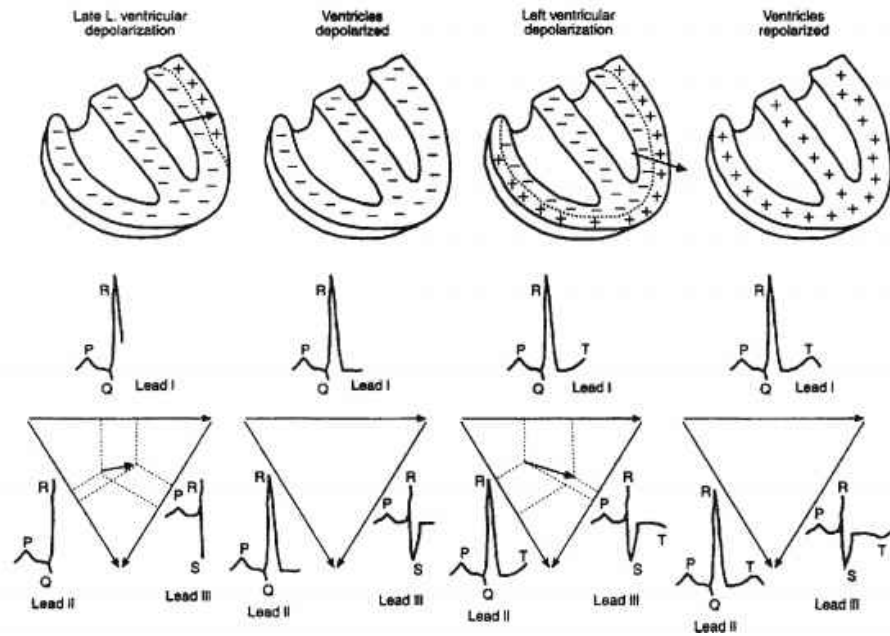


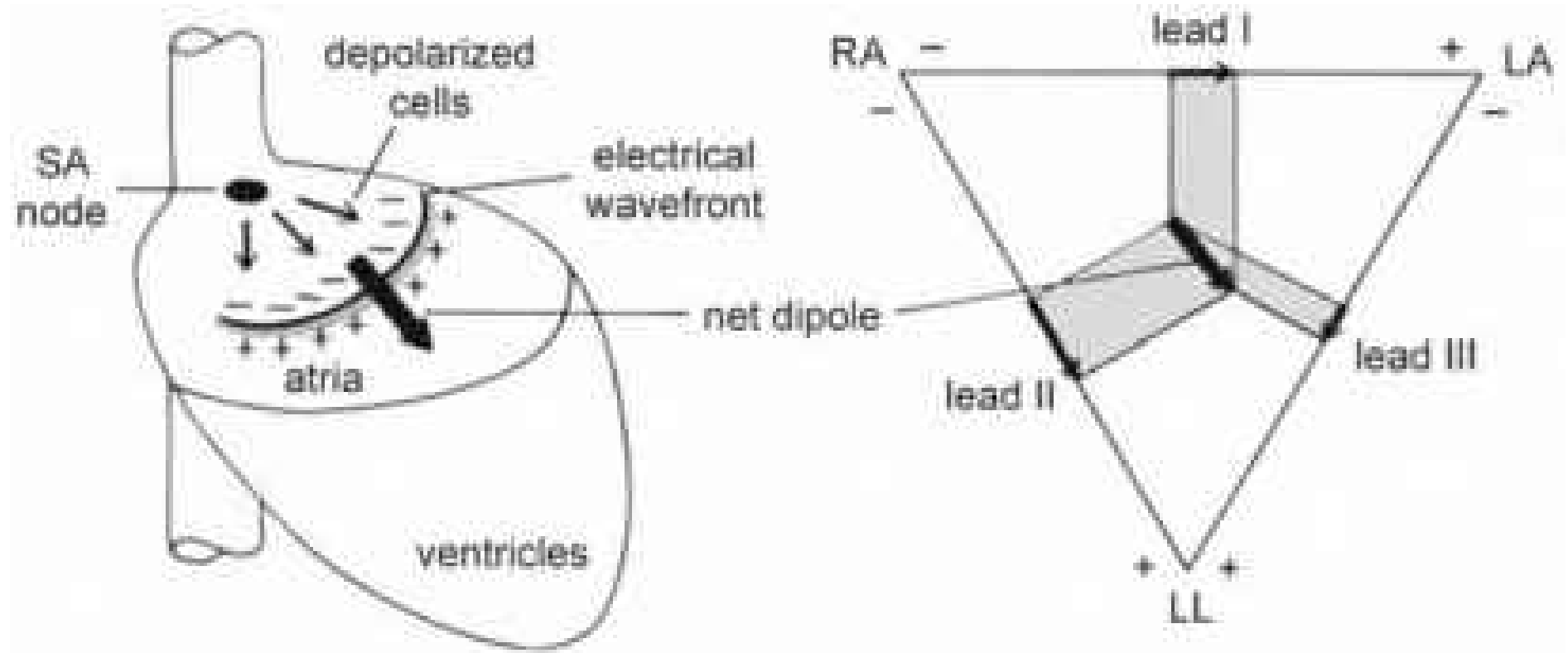


Progression of depolarization

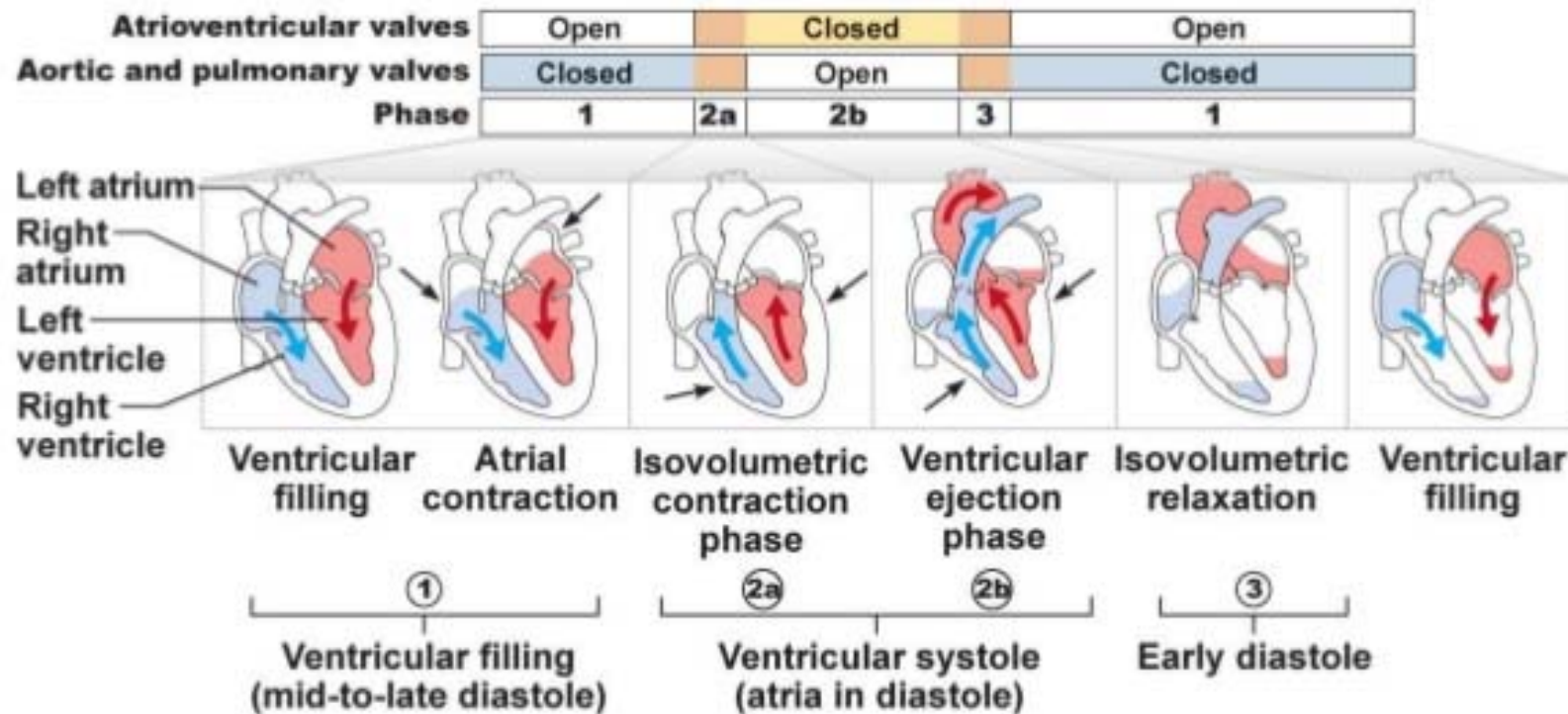


End of depolarization and repolarization

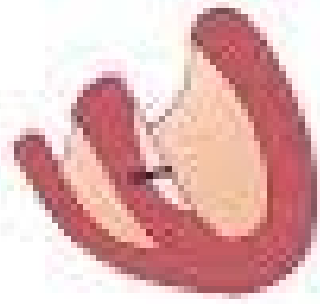




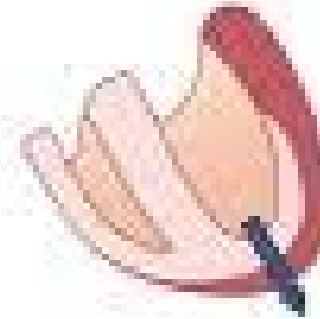
Steps of the Cardiac Cycle



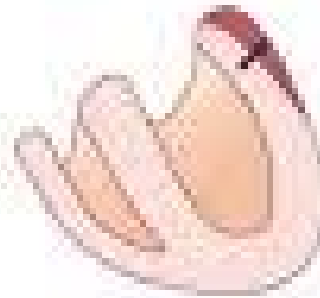
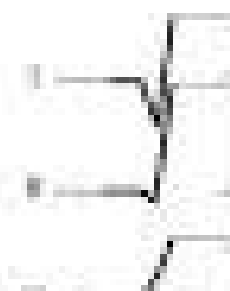
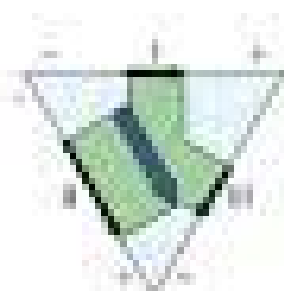
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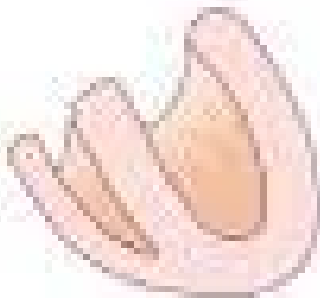
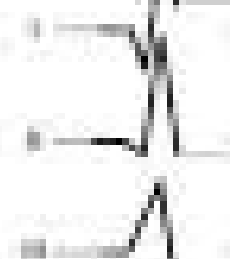
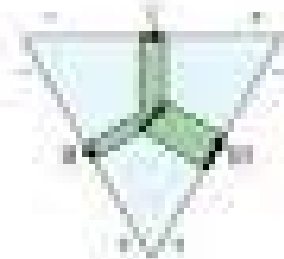
I
normal



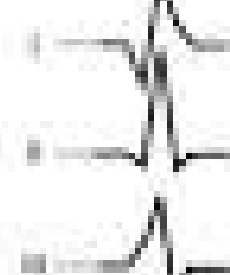
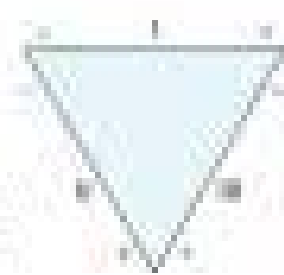
II
normal

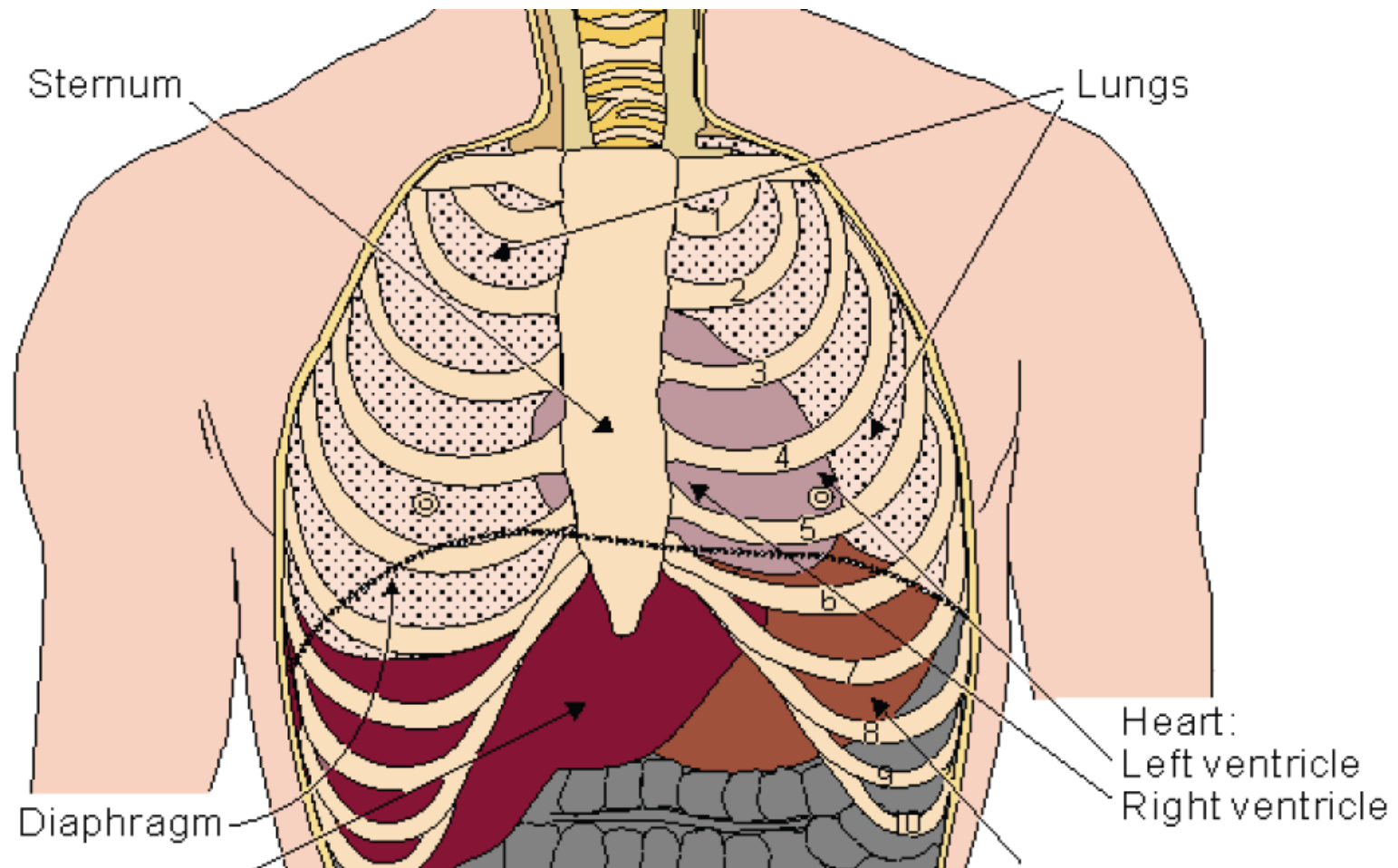


III
normal

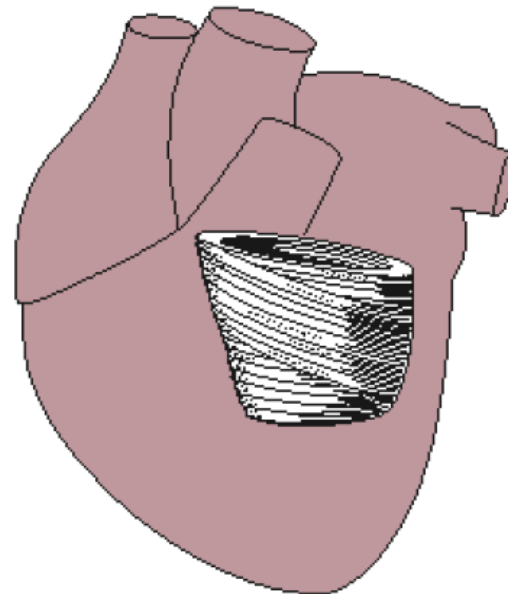
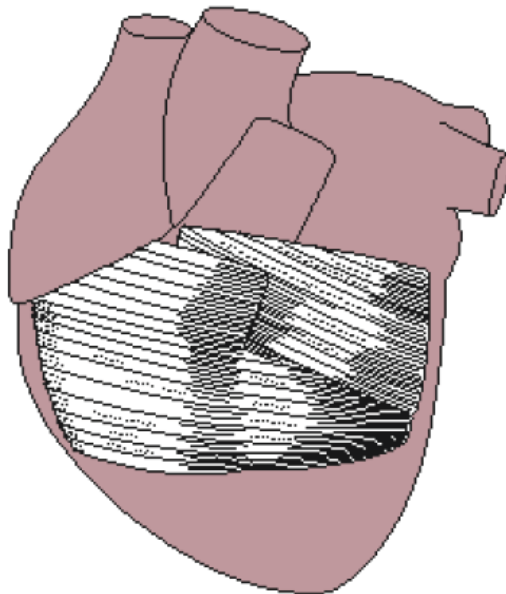
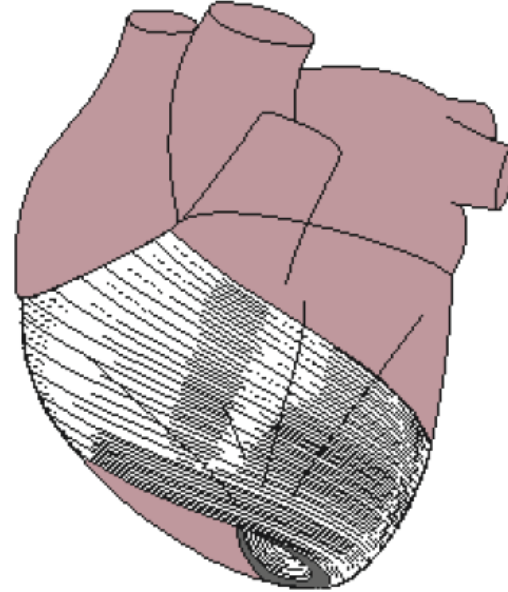
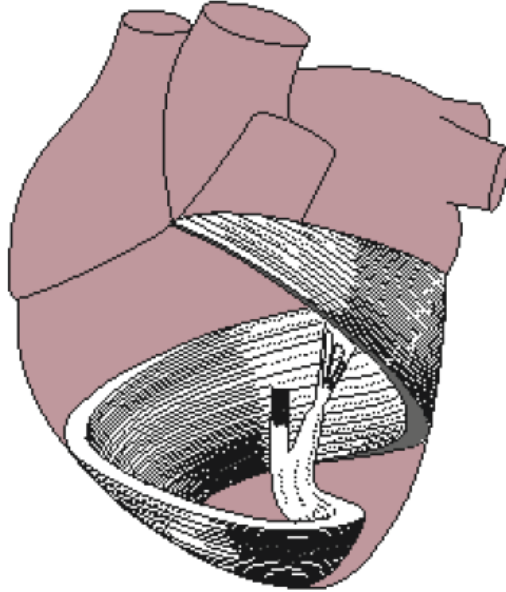


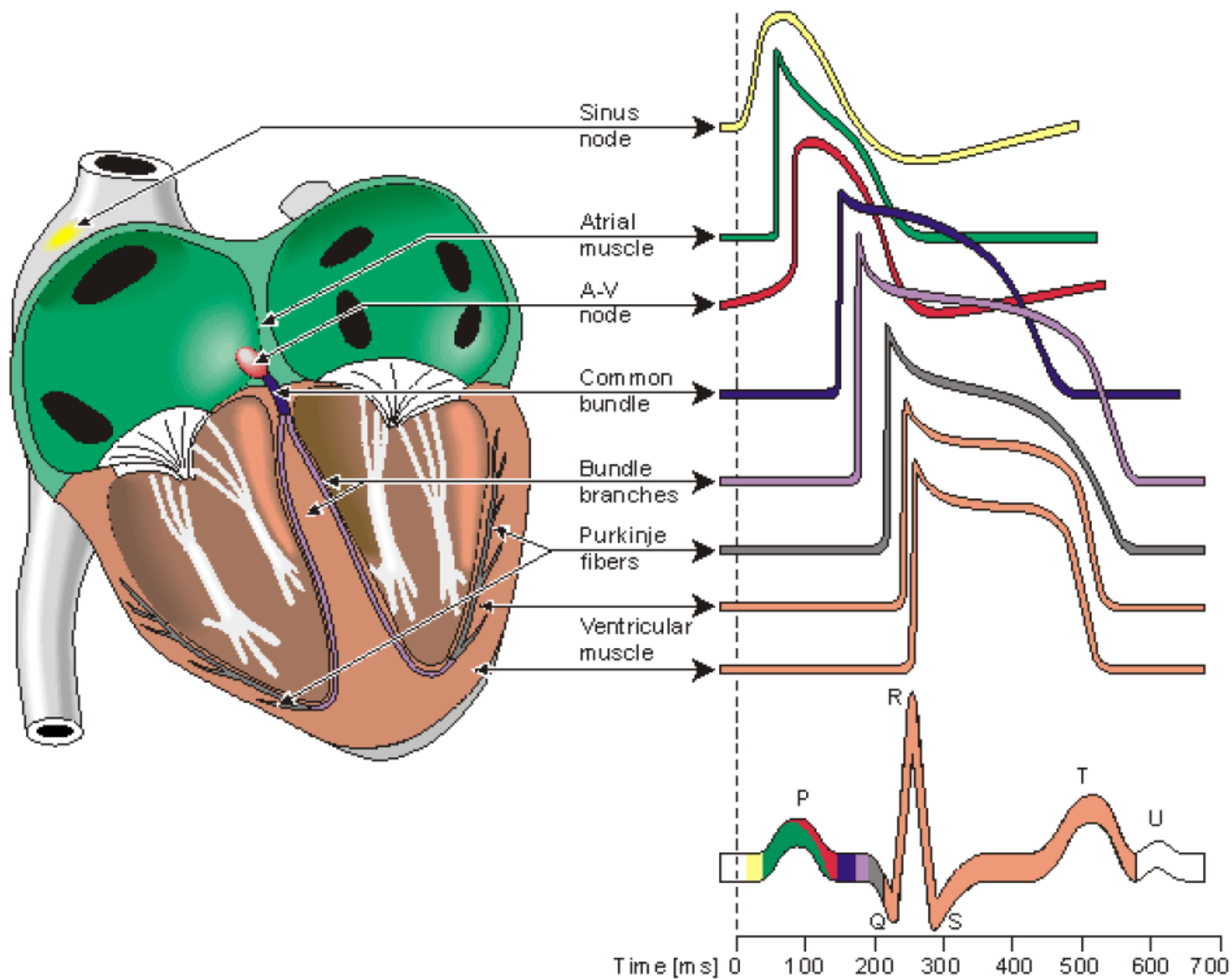
IV
septated

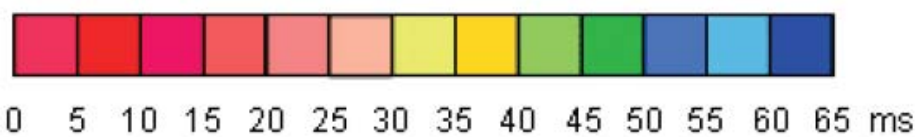
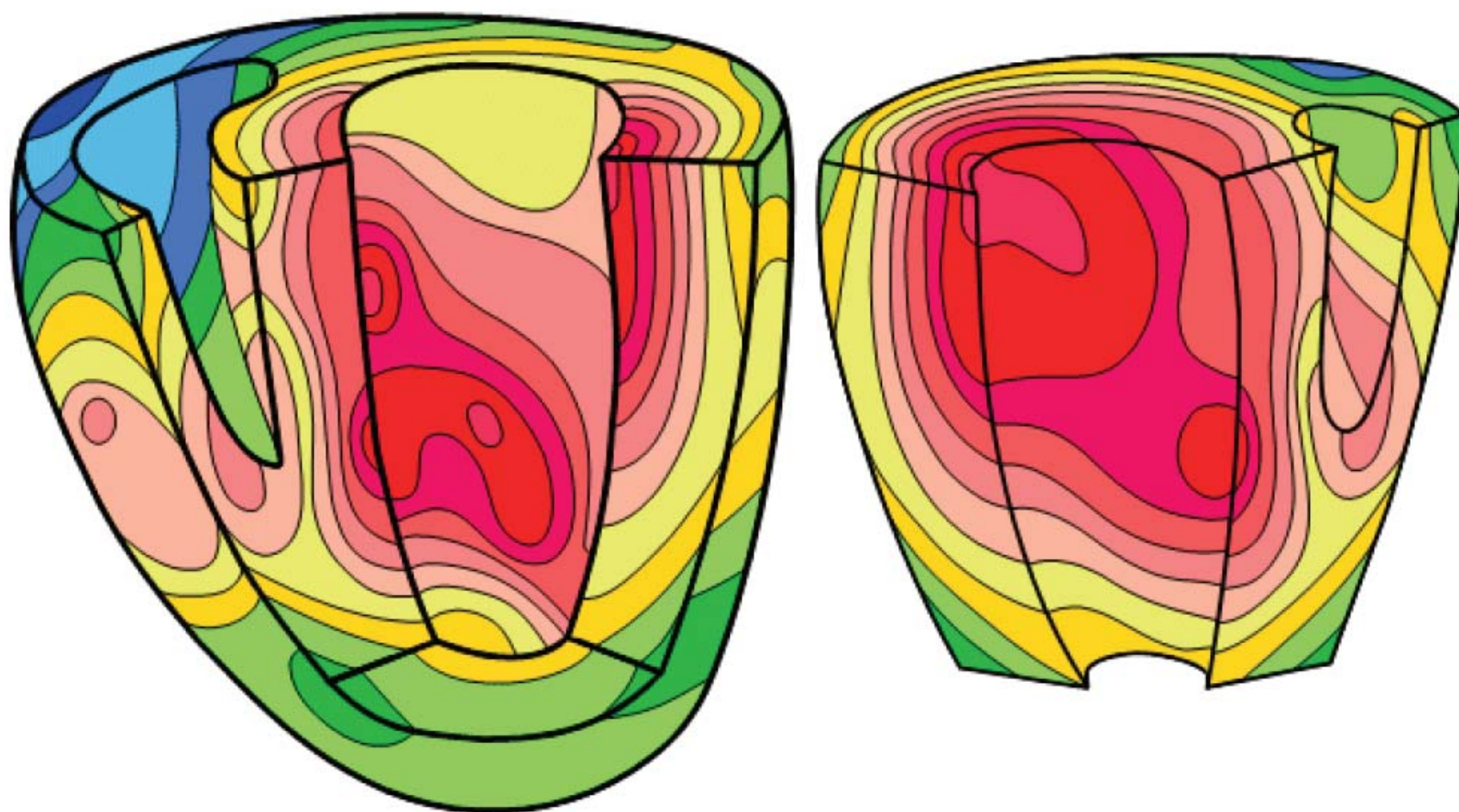


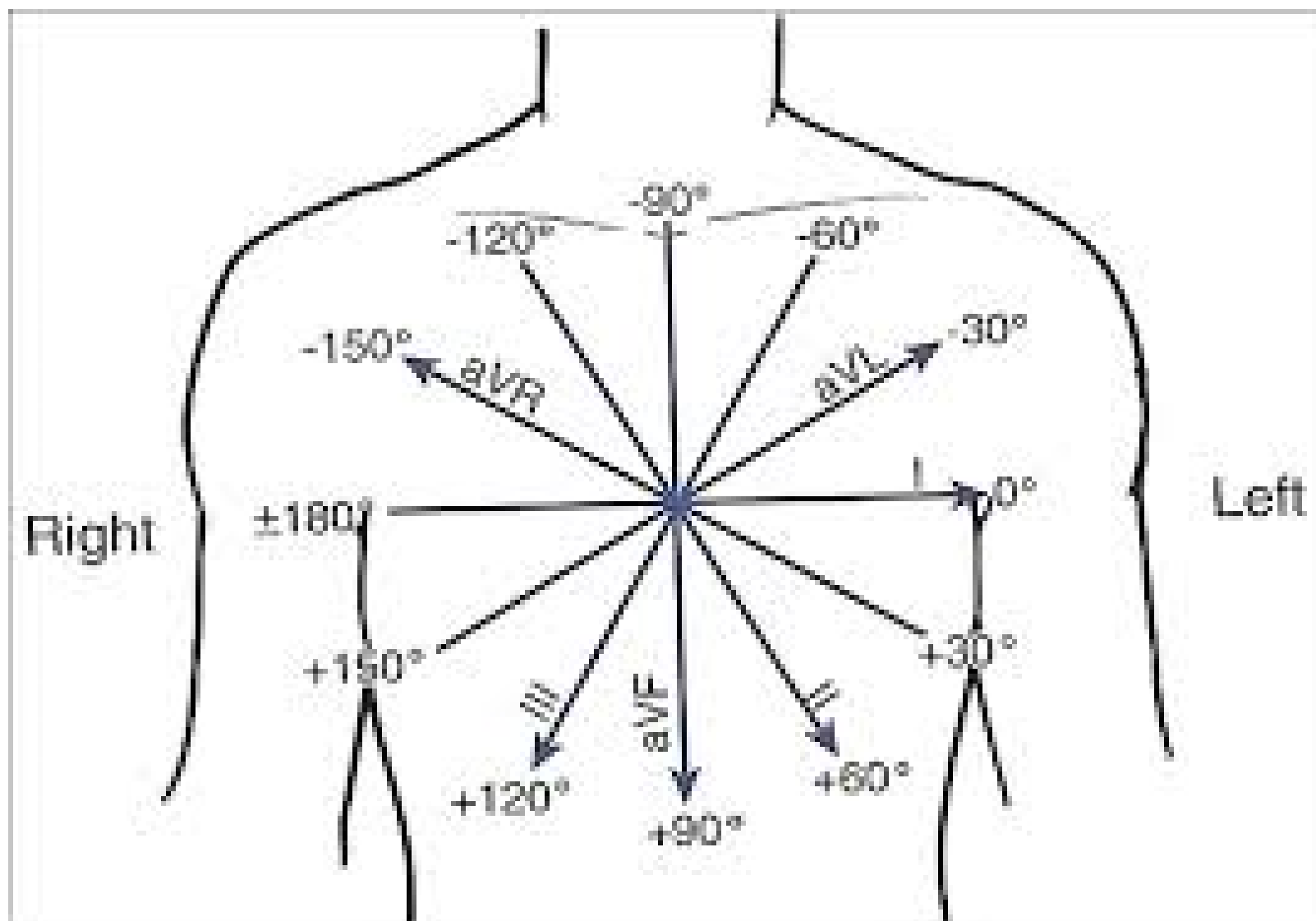


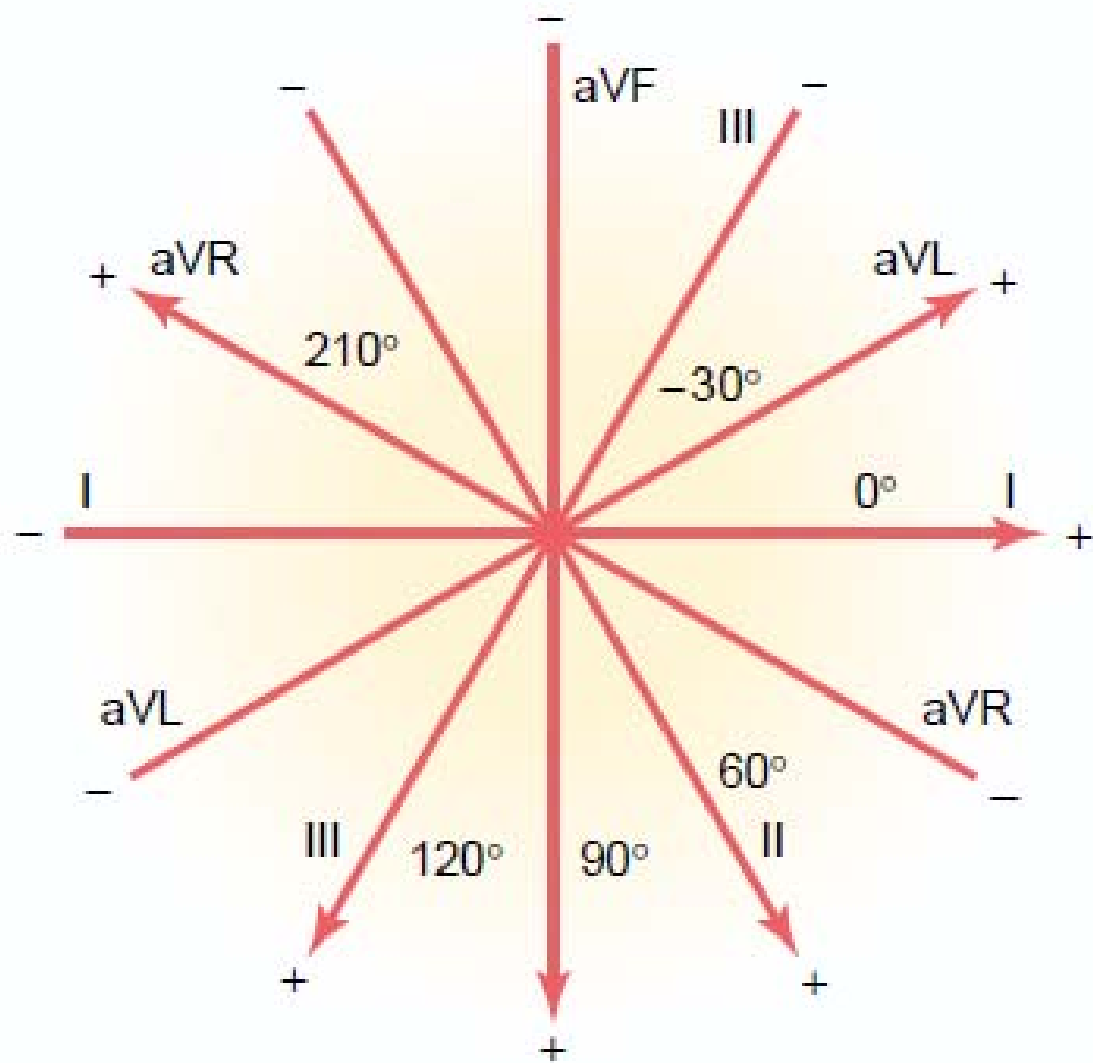
6. The Heart

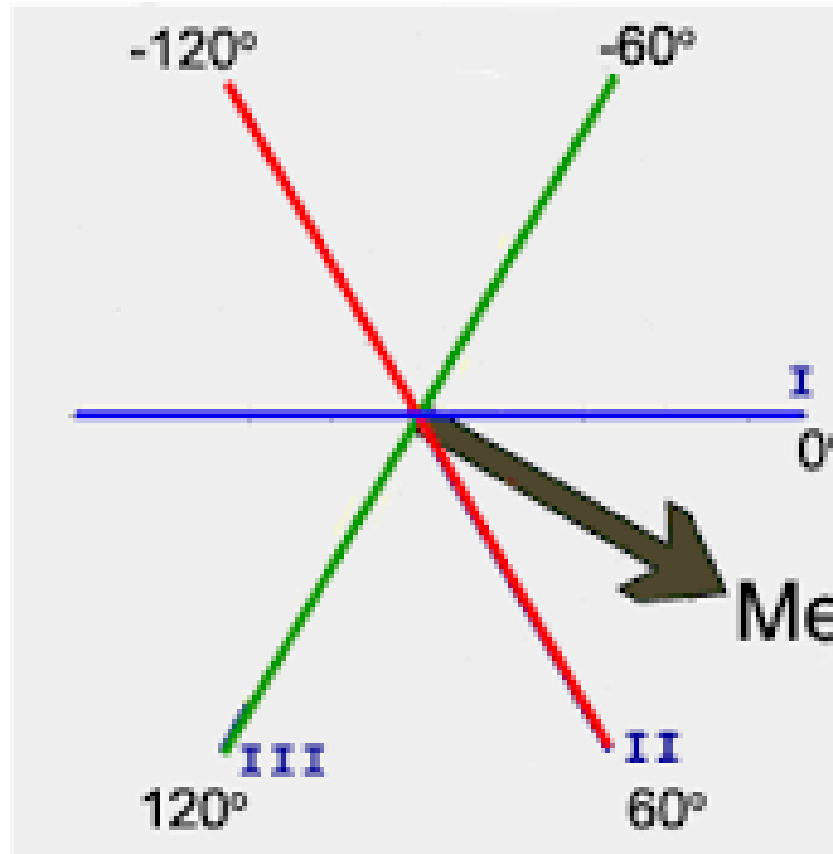










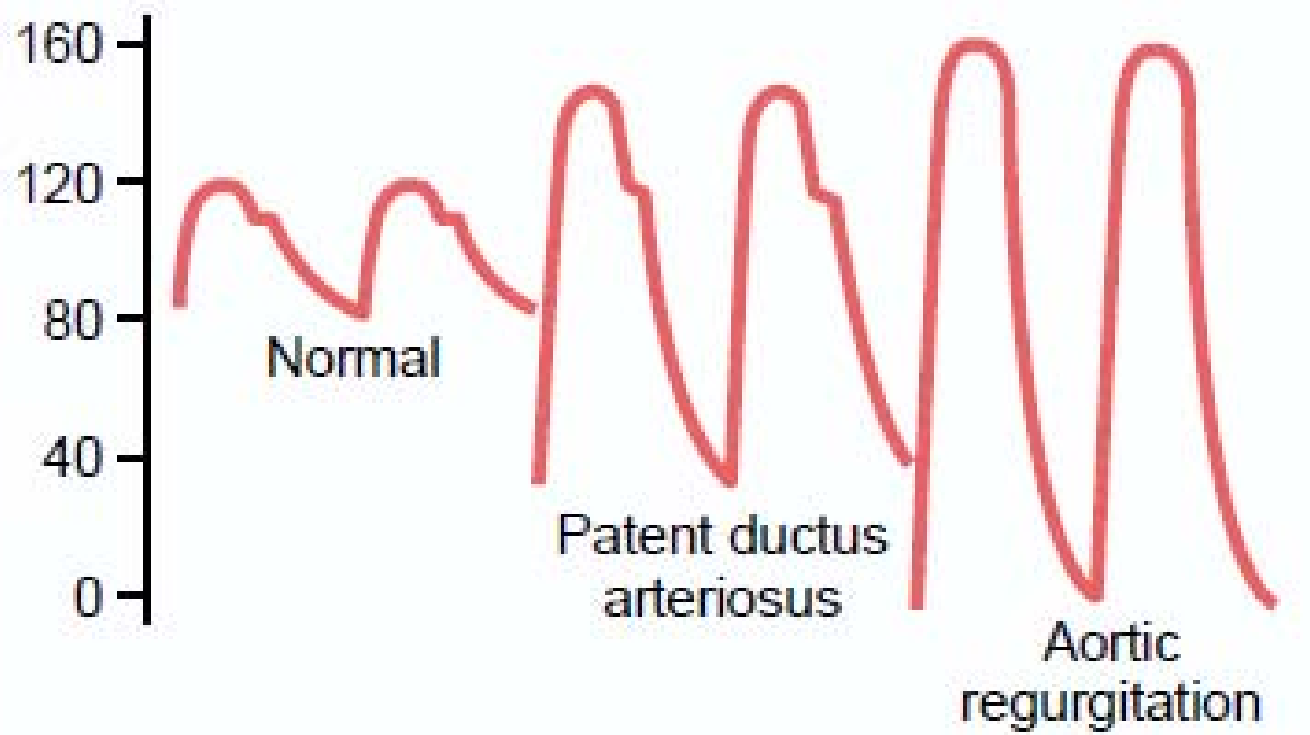
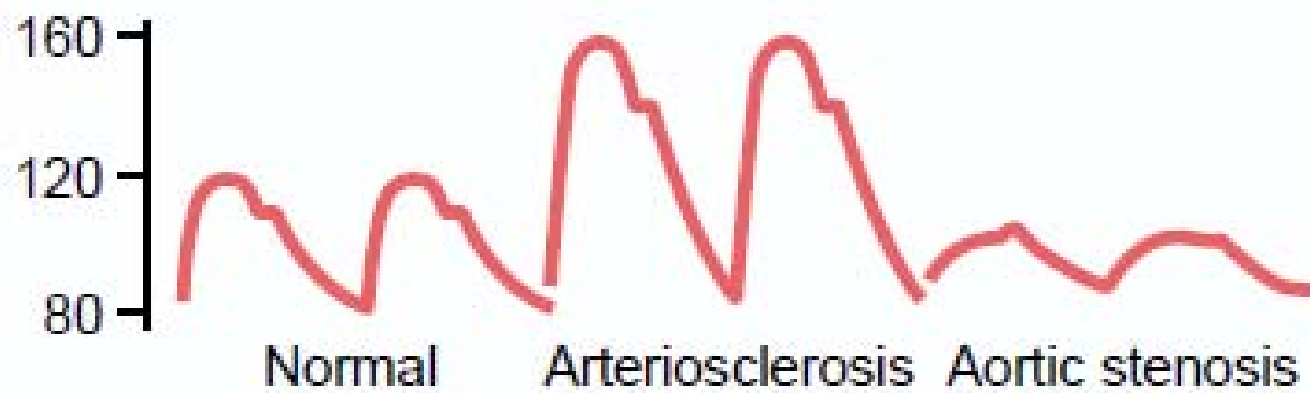


Arterial Blood Pressure

- Systolic
- Diastolic
- Pulse Pressure
- Mean pressure

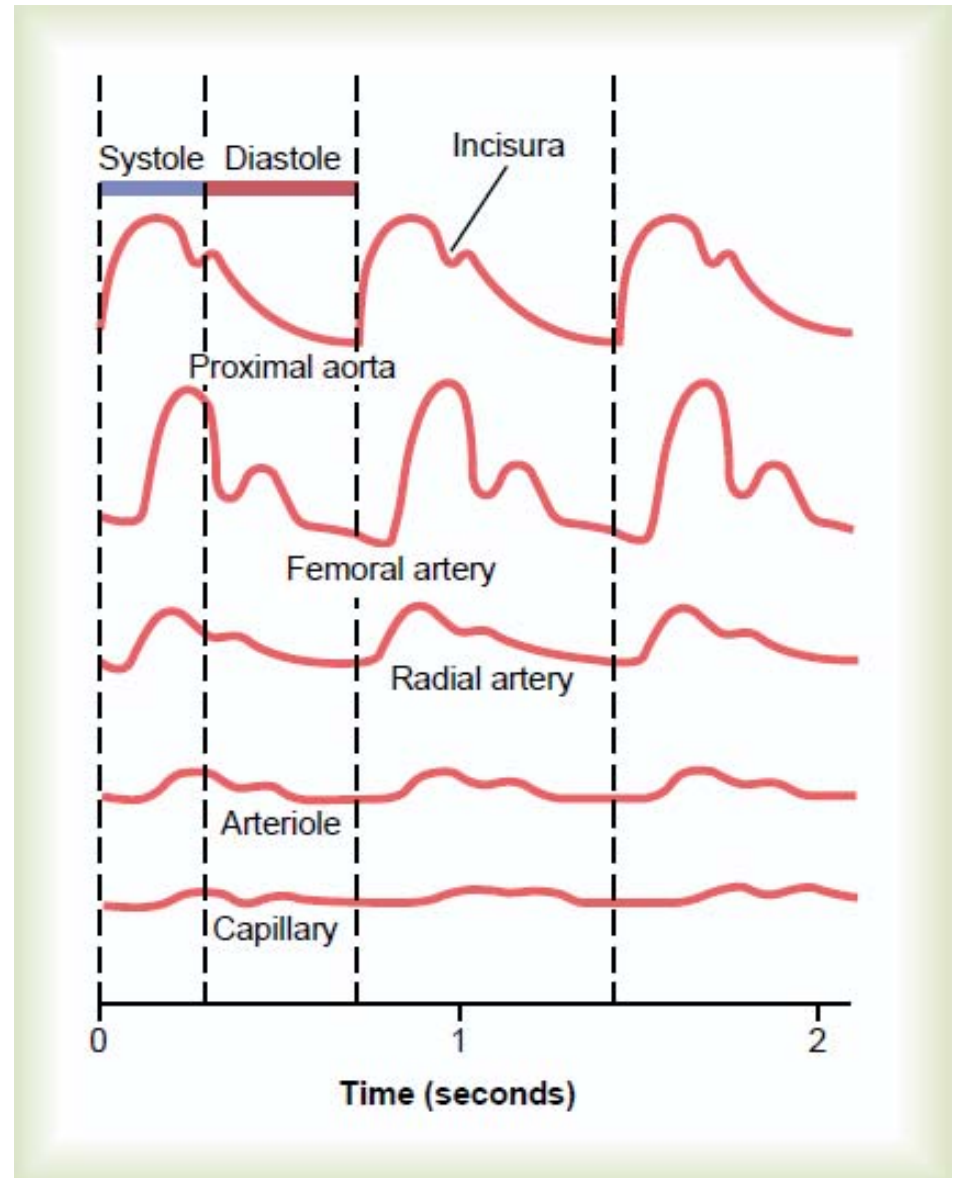
Pulse pressure

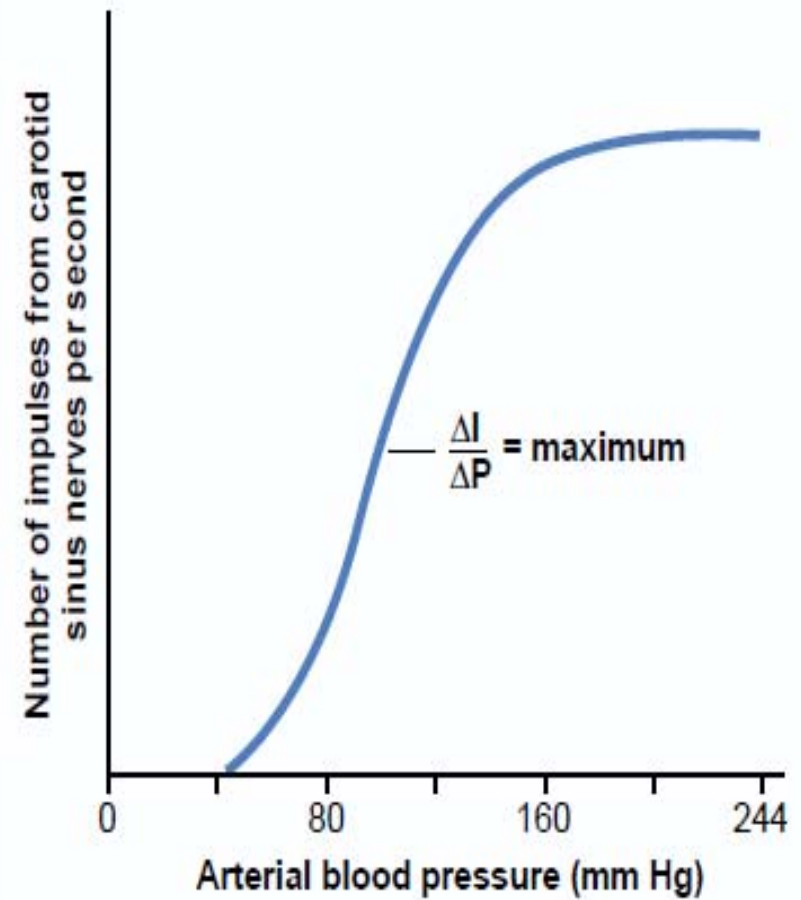
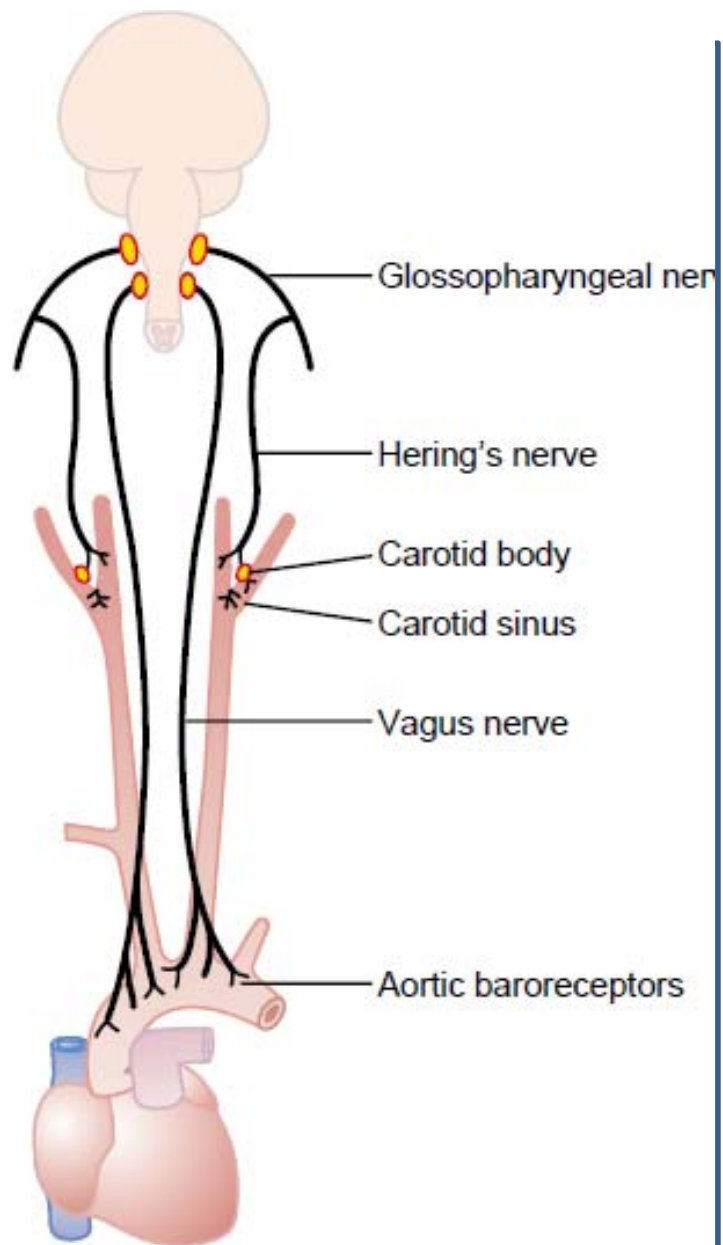
- Determinants
 - Stroke volume output
 - Arterial compliance
 - Character of ejection during systole
- Ratio of stroke volume and arterial compliance

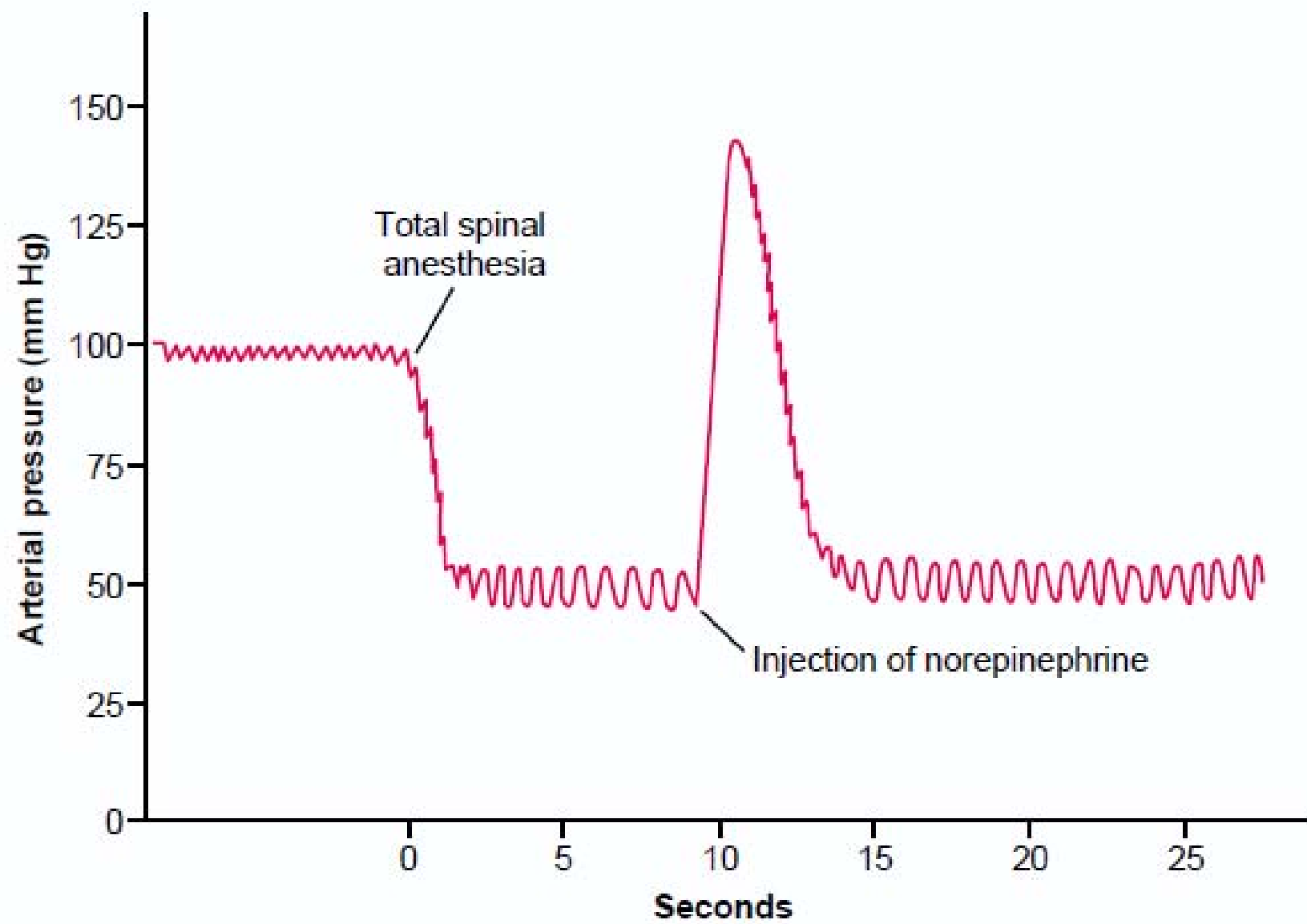


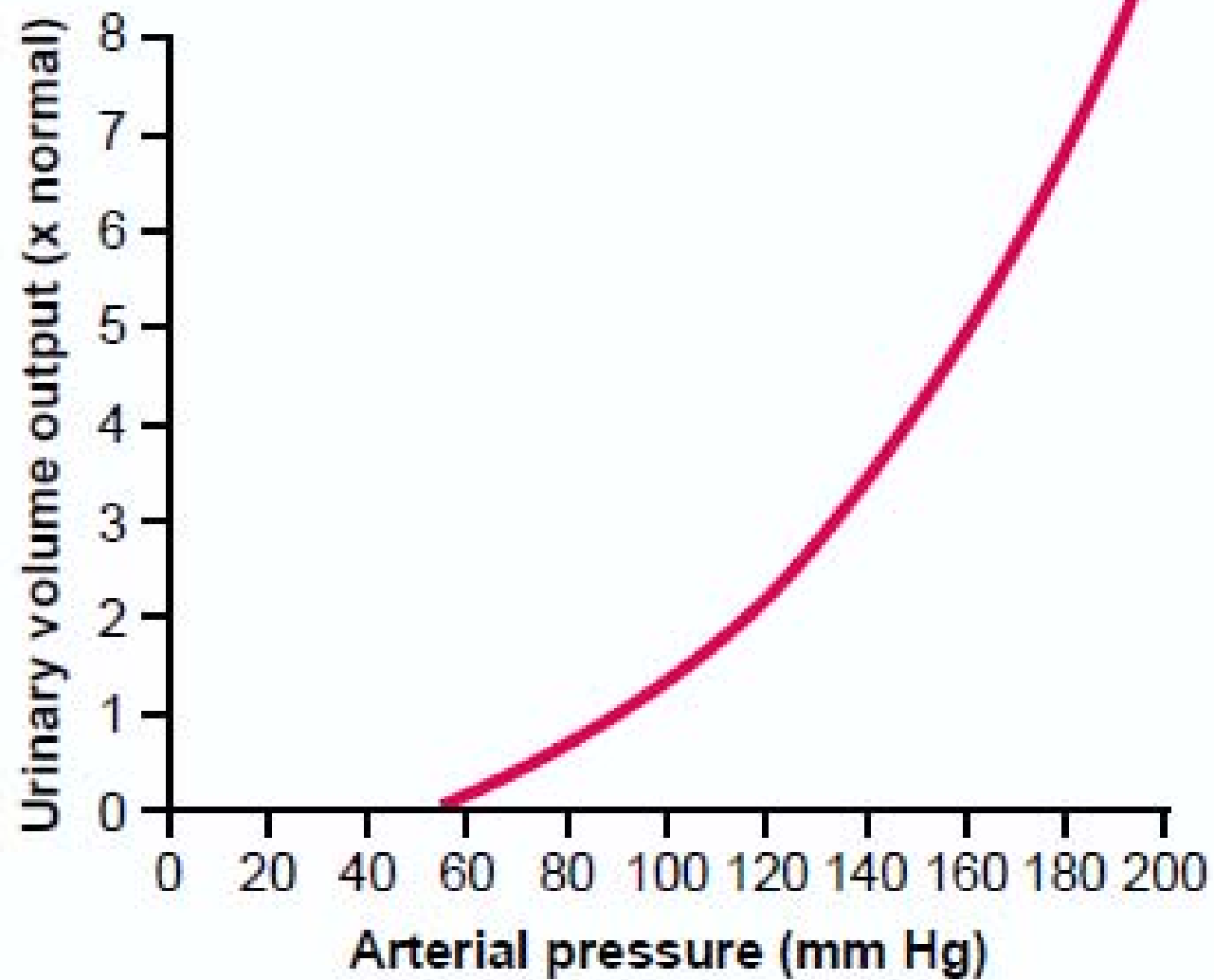
Damping of the pressure pulse

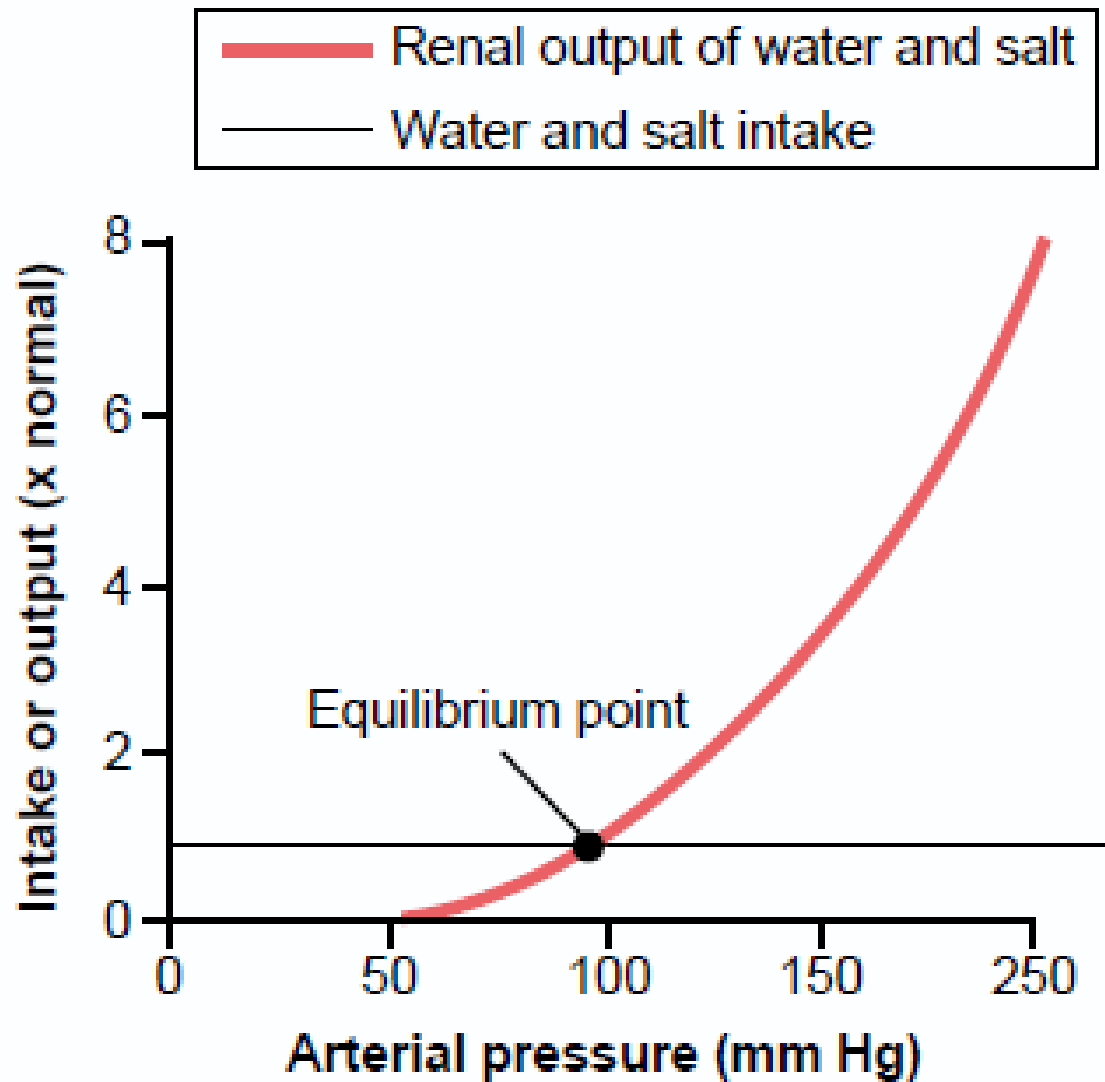
Due to resistance and compliance of blood vessel

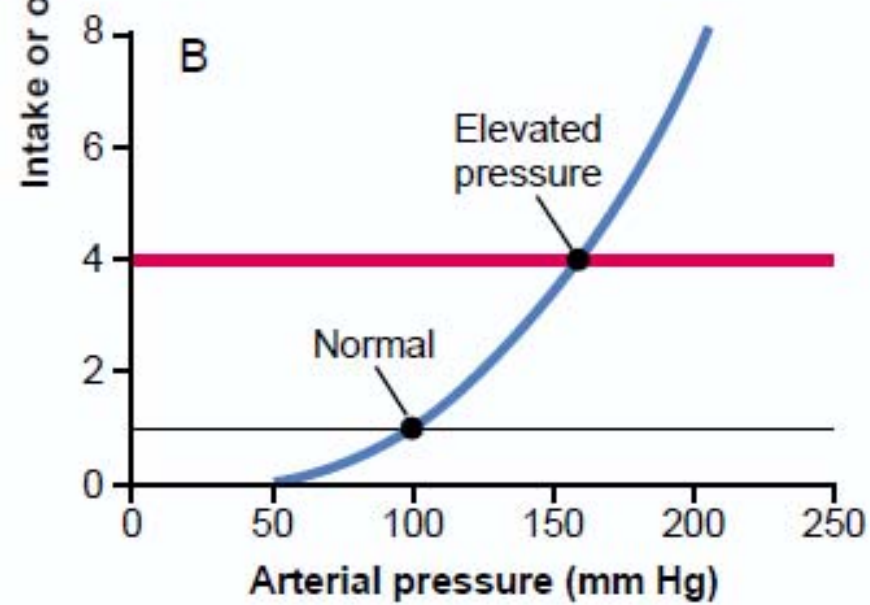
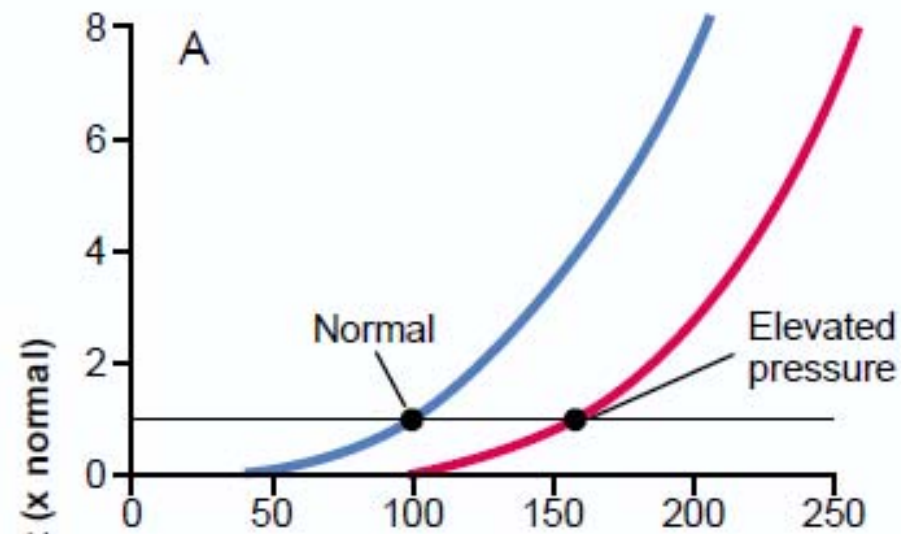


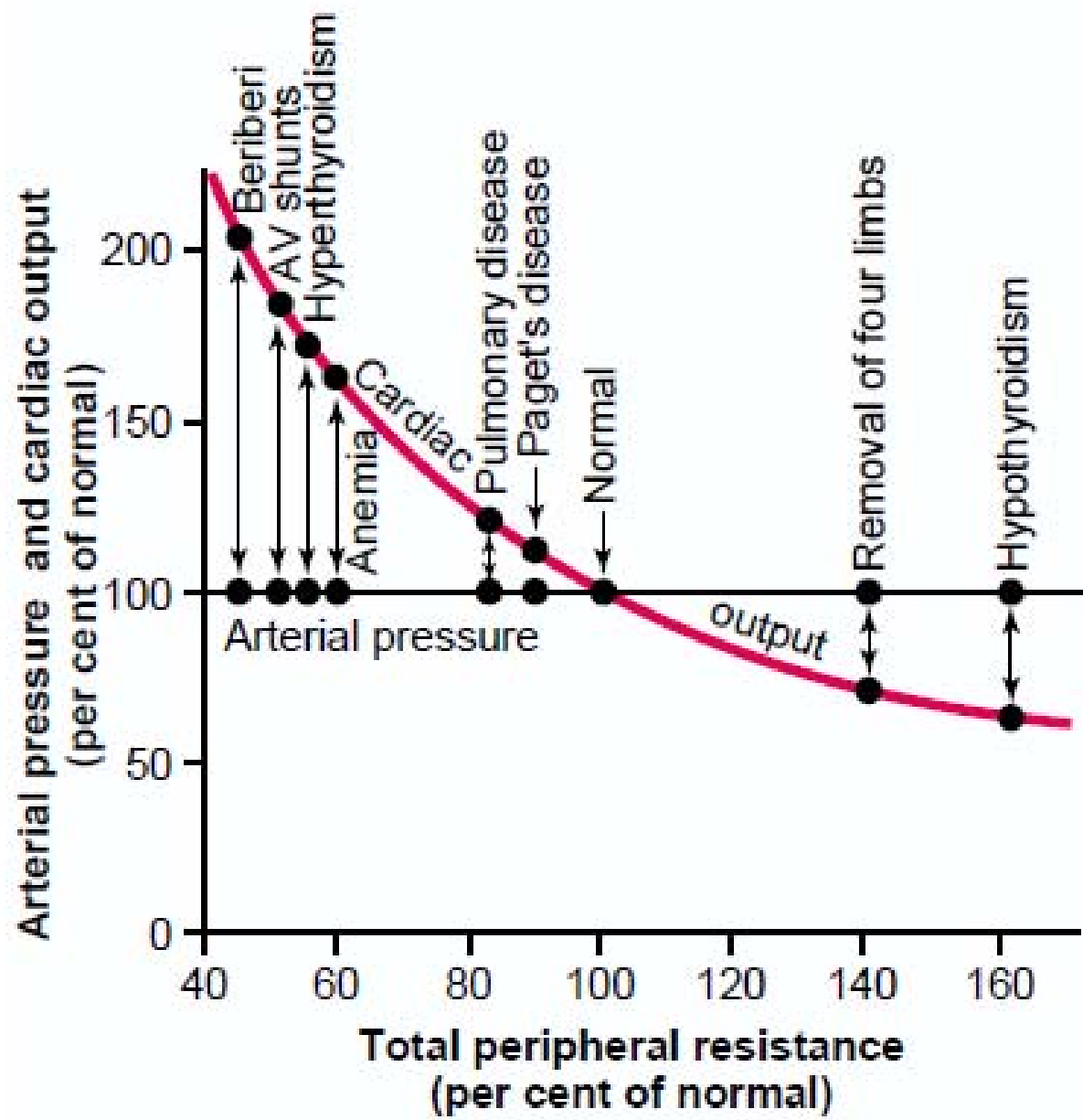




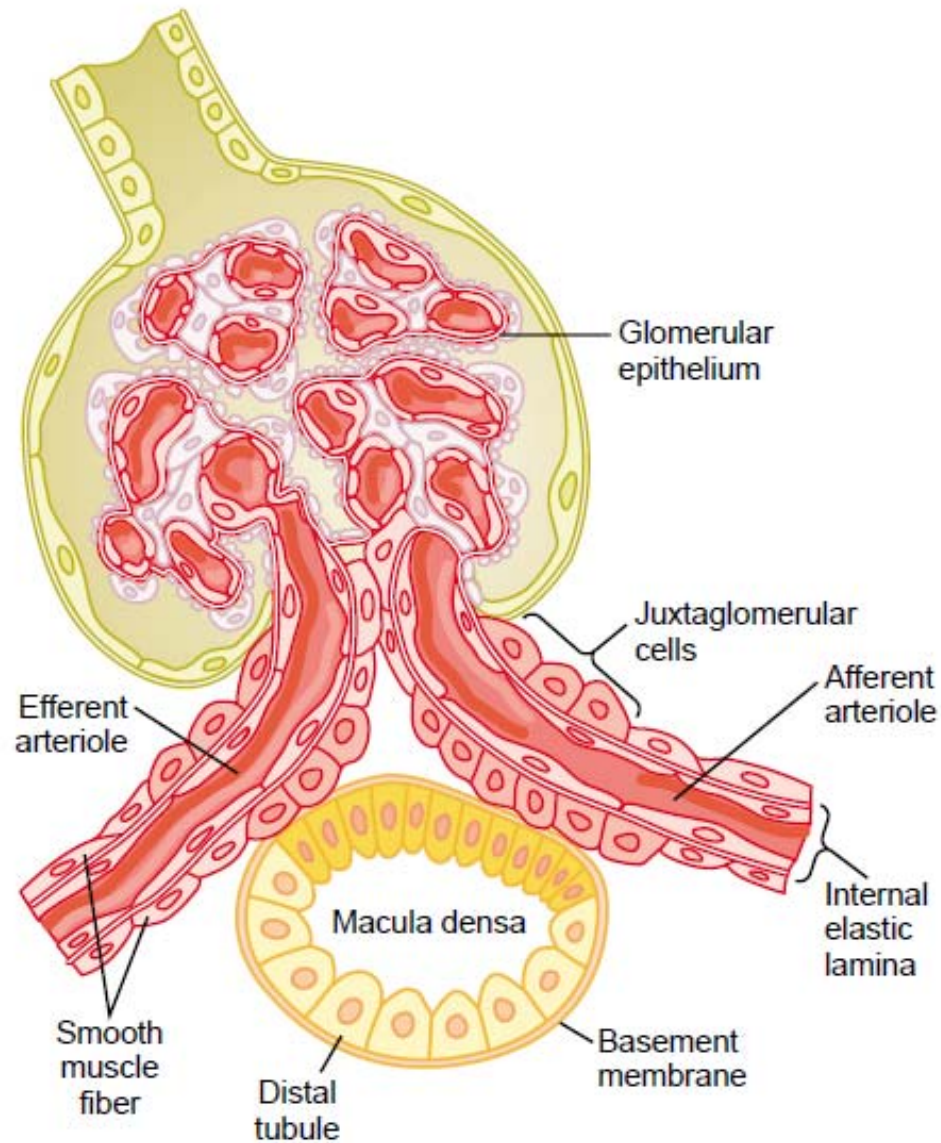


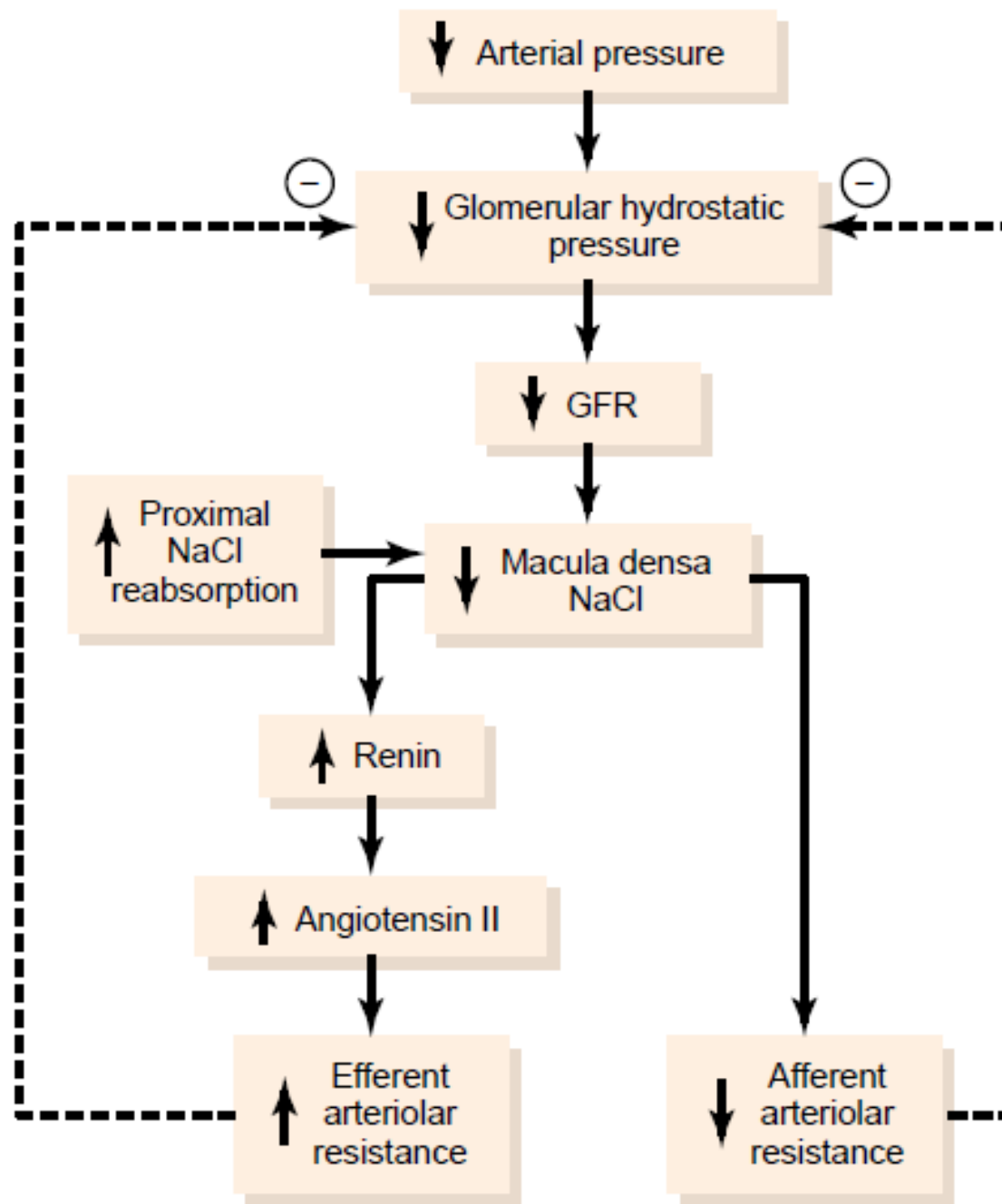


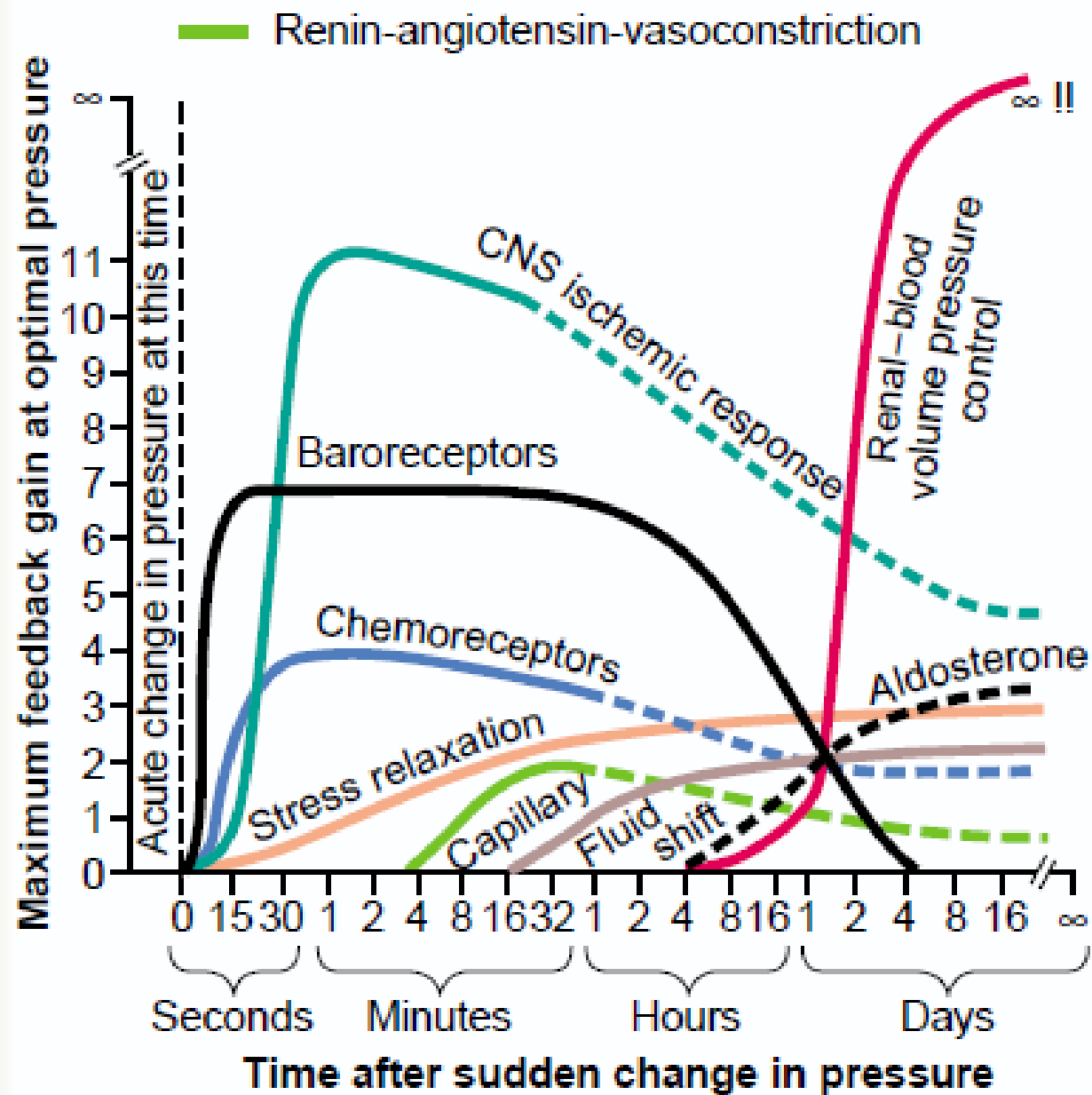




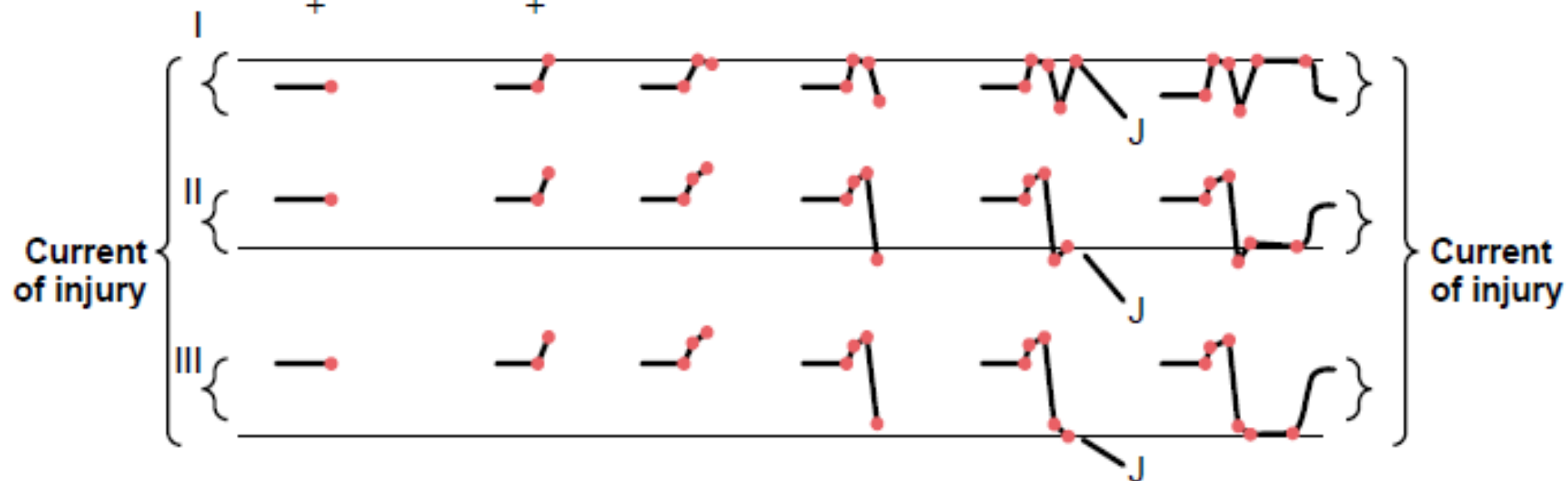
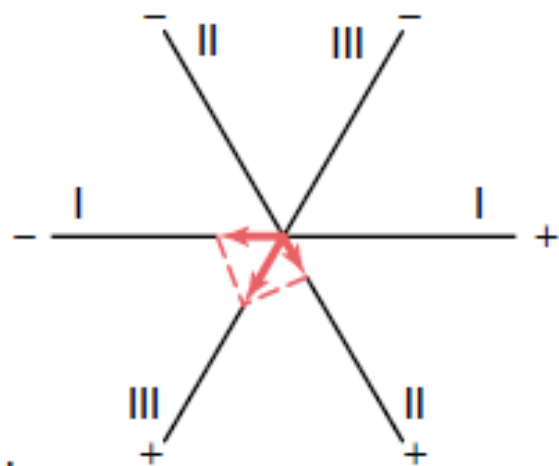
Renin- angiotensin System

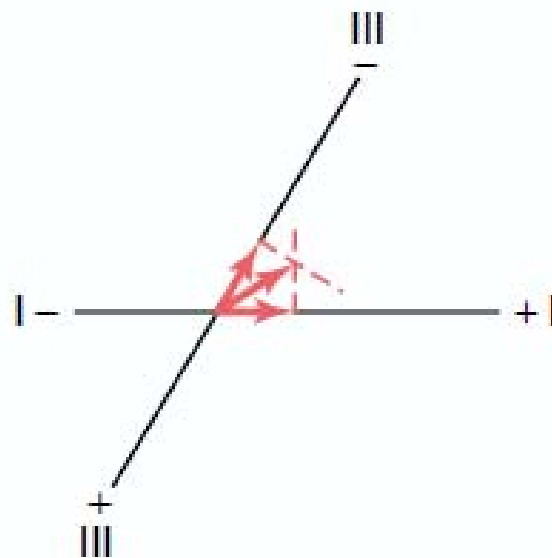
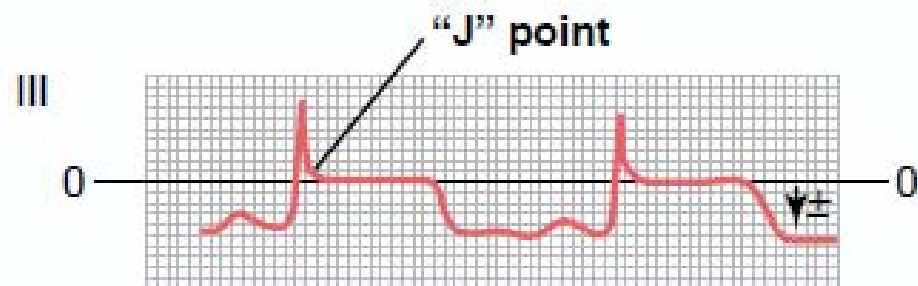
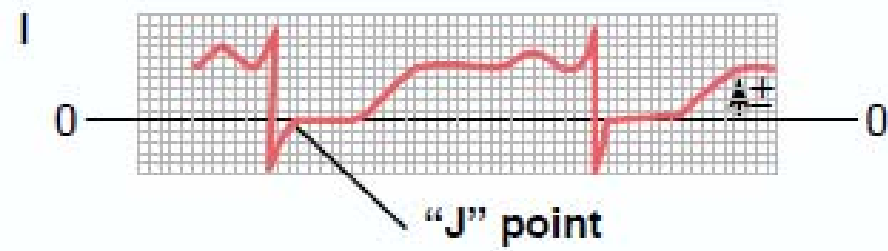






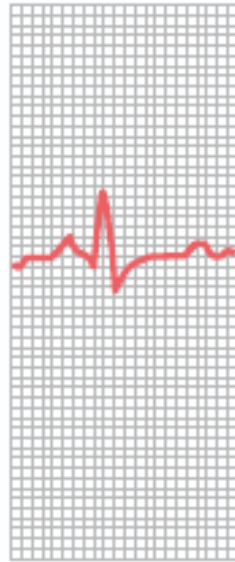
Injured area







I



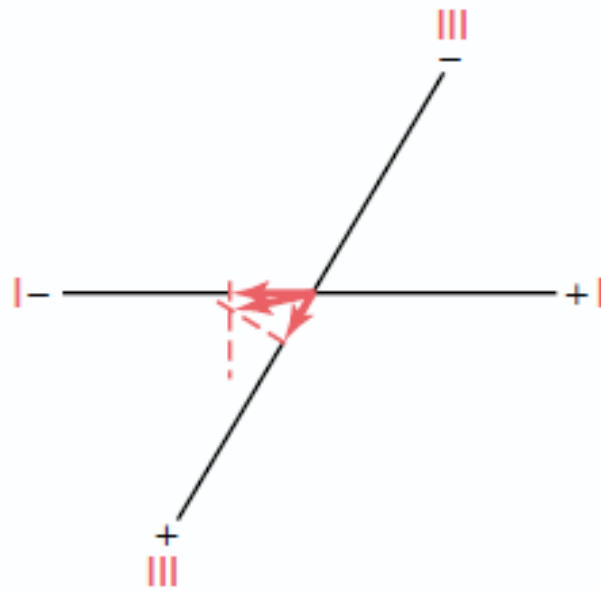
II



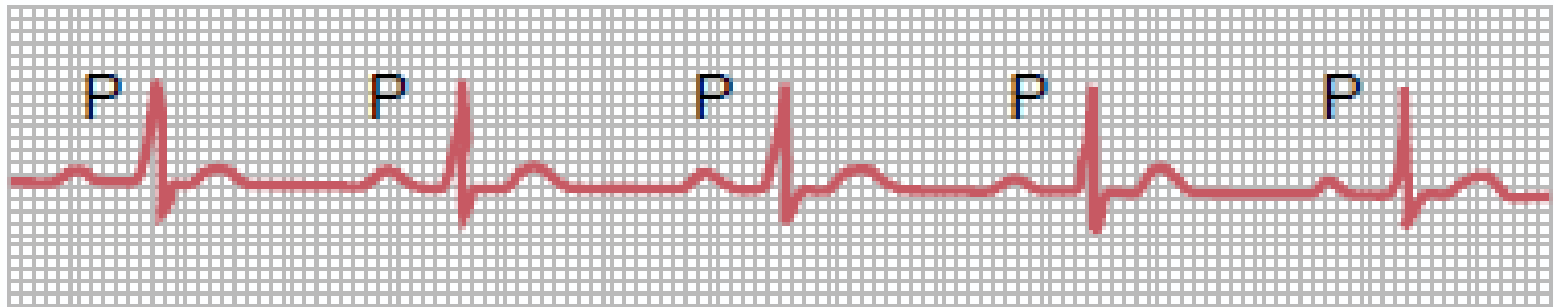
III



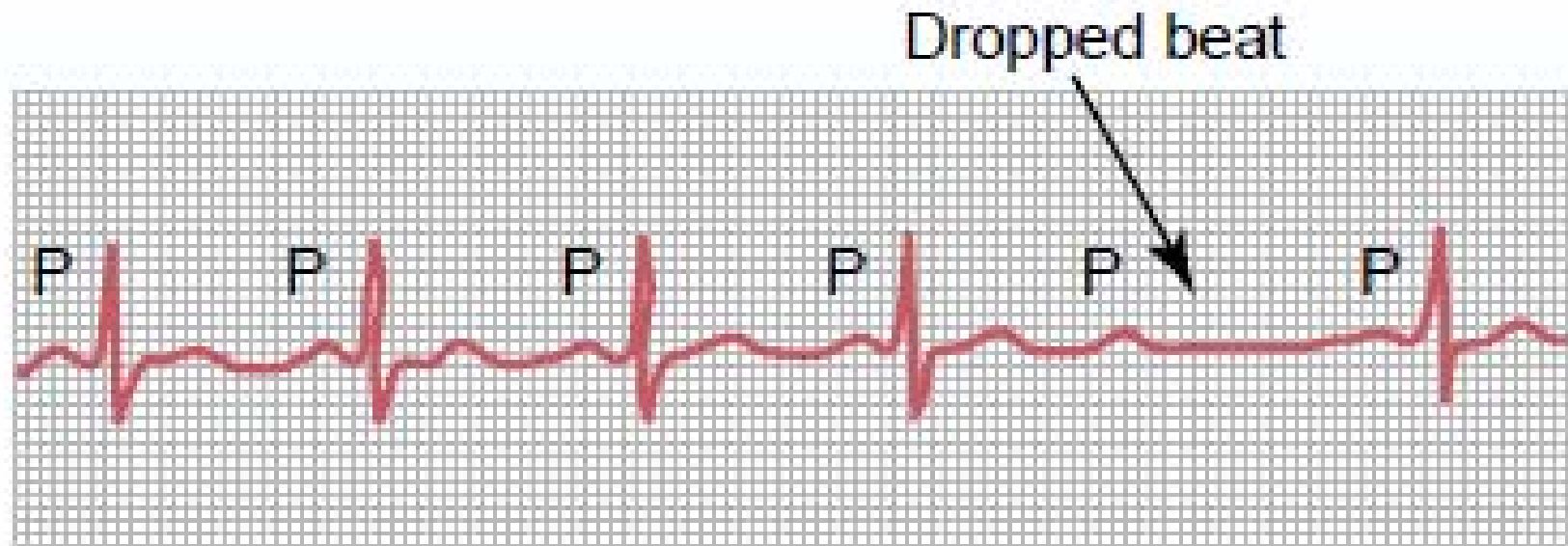
V₂



1st degree heart block



2nd degree heart block



Complete heart block

