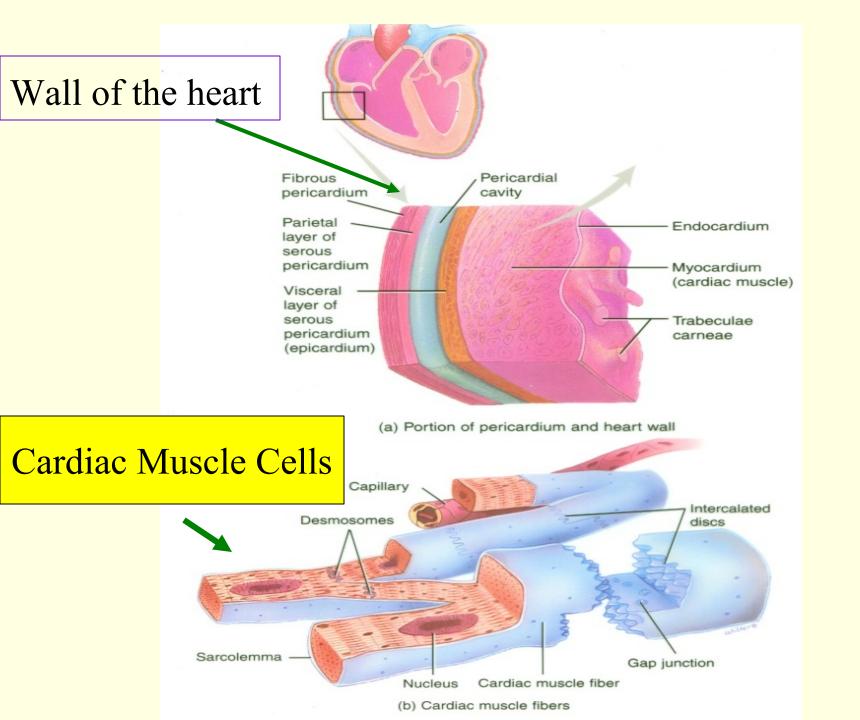
Cardiac Muscle Physiology

Faisal Mohammed, MD, PhD Alaa Bawaneh, MD.PhD

Objectives:

By The end of this lecture students should be able to:

- Distinguish the cardiac muscle cell microstructure
- Describe cardiac muscle action potential
- Point out the functional importance of the action potential
- Outline the intracellular calcium homeostasis



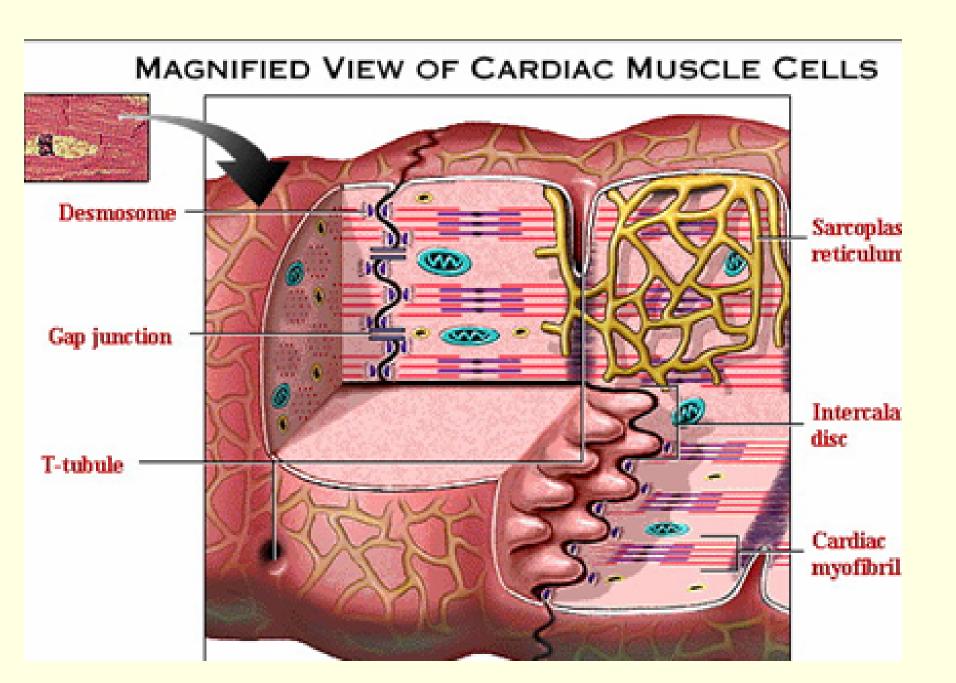
Layers of the Heart Wall

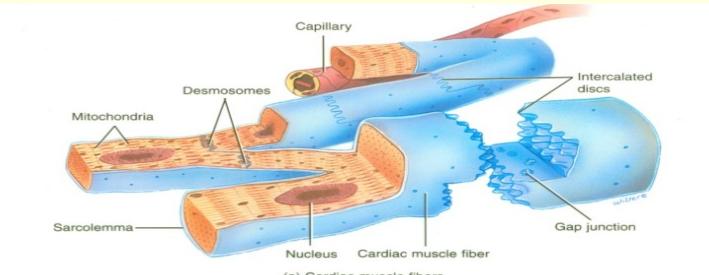
- 1. Epicardium (external layer)...prevent the heart from overstretching as we will see later when we discuss Frank-Starling law of the heart.
 - Visceral layer of serous pericardium
 - Smooth, slippery texture to outermost surface
- 2. Myocardium
 - 95% of heart is cardiac muscle
- 3. Endocardium (inner layer)
 - Smooth lining for chambers of heart, valves and continuous with lining of large blood vessels

Cardiac Muscle Tissue and the Cardiac Conduction System

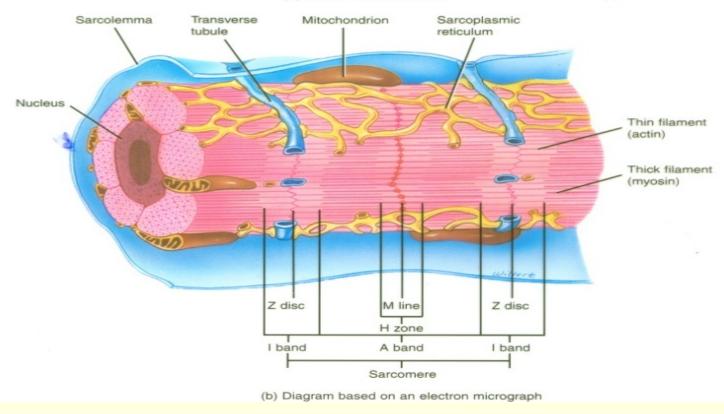
Histology

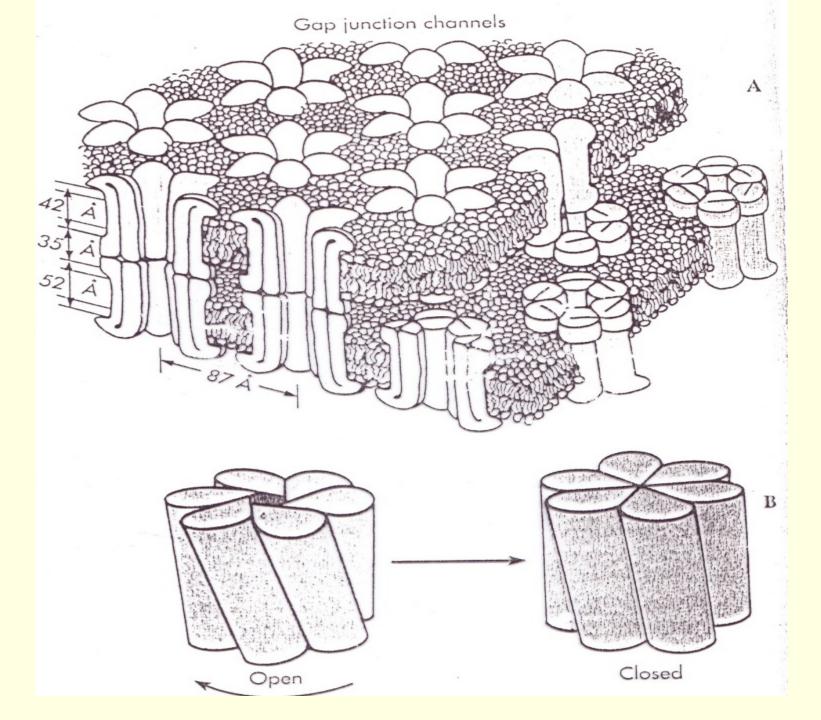
- Shorter and less circular than skeletal muscle fibers
- Branching gives "stair-step" appearance
- Usually, one centrally located nucleus
- Ends of fibers connected by intercalated discs
- Discs contain desmosomes (hold fibers together) and gap junctions (allow action potential conduction from one fiber to the next) → syncytium
- Mitochondria are larger and more numerous than skeletal muscle
- Same arrangement of actin and myosin



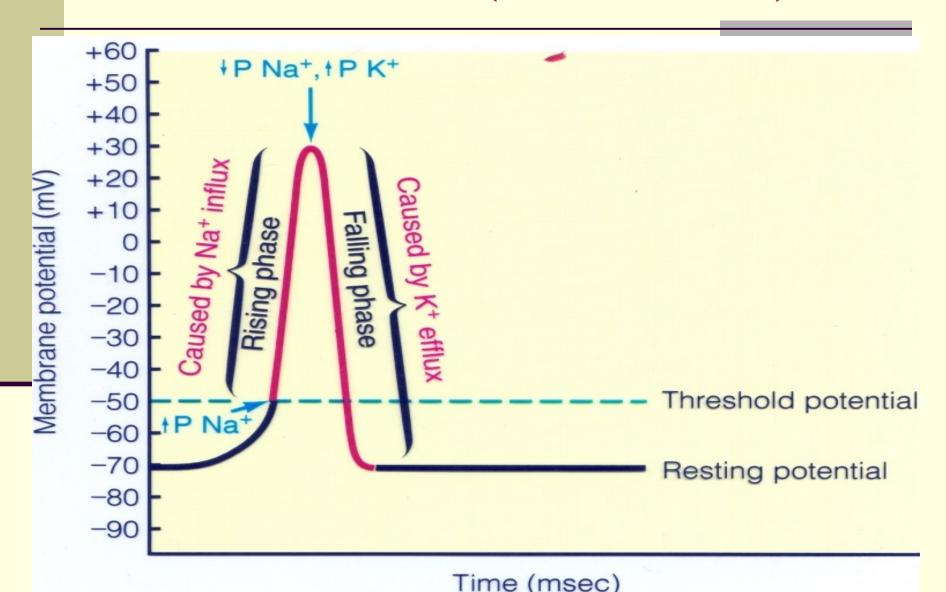


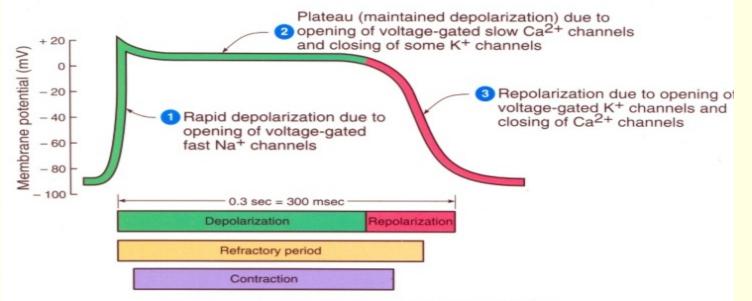
(a) Cardiac muscle fibers



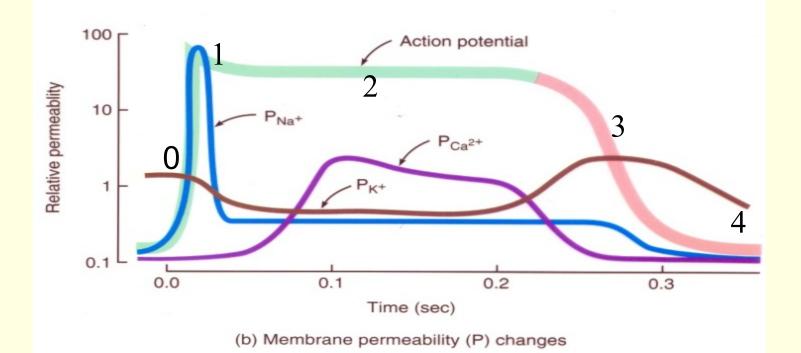


Permeability Changes and Ionic Fluxes During an Action Potential (skeletal Muscle)

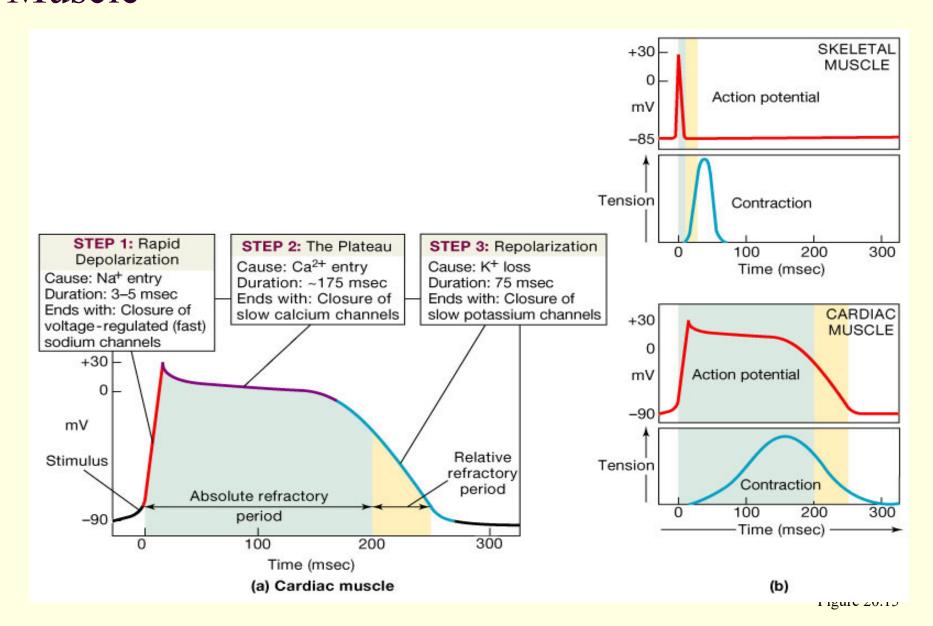




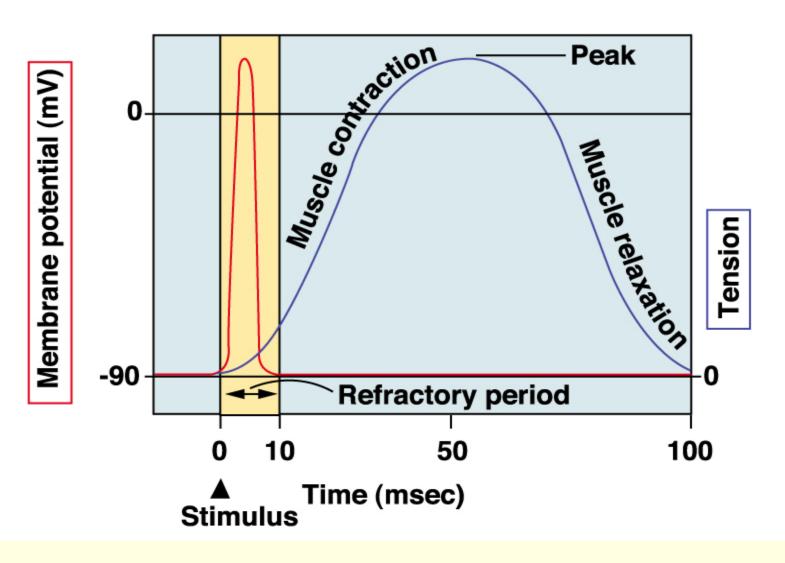
(a) Action potential, refractory period, and contraction



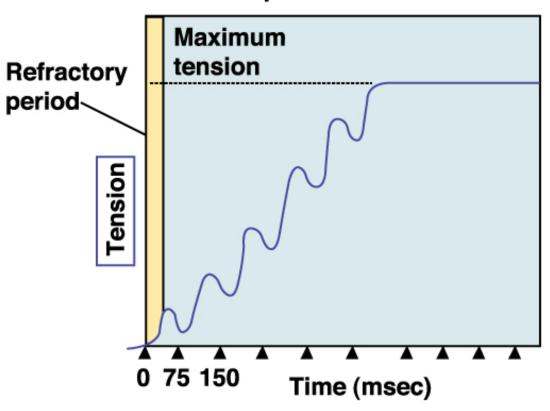
The Action Potential in Skeletal and Cardiac Muscle



Skeletal muscle fast-twitch fiber

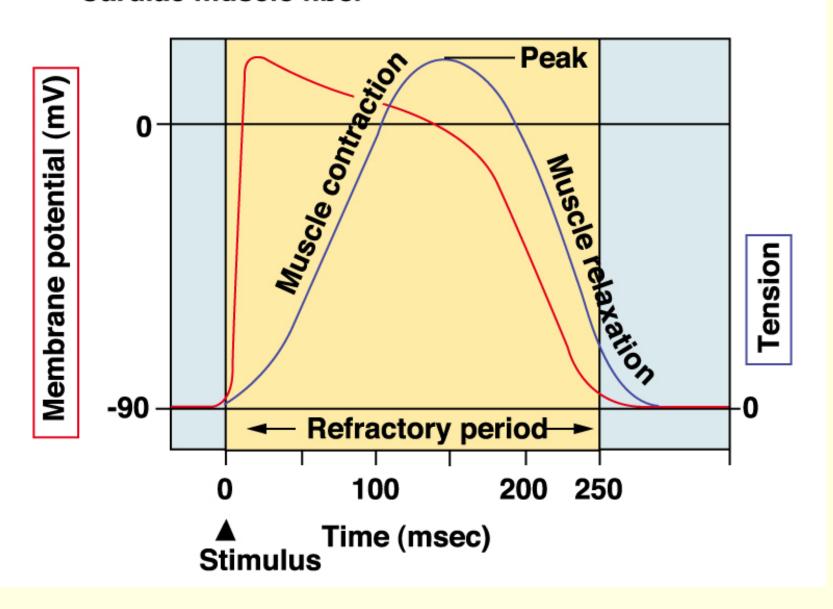


Tetanus in a skeletal muscle. Action potentials not shown.

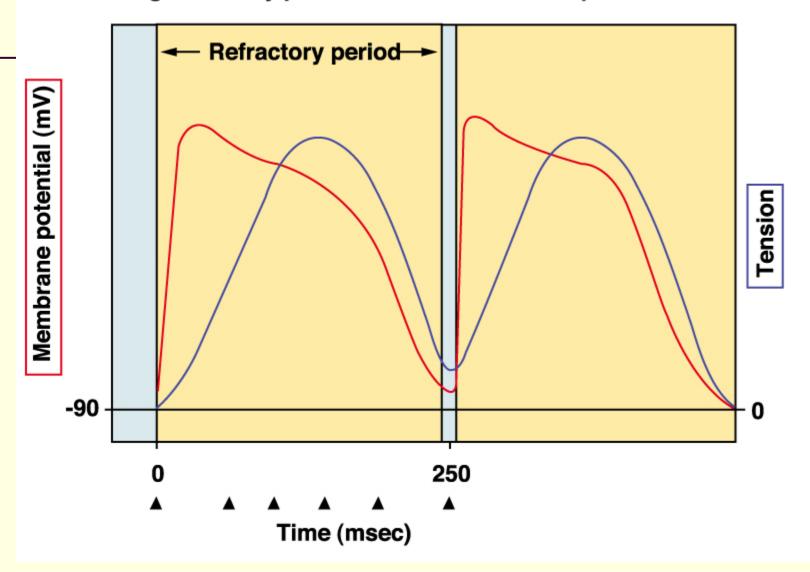


▲ = Stimulus for action potential

Cardiac muscle fiber



Long refractory period in a cardiac muscle prevents tetanus.



Cardiac and Skeletal Muscles Differences

Skeletal muscle

- Neurogenic

 (motor neuron-end plate-acetylcholine)
- Insulated from each other
- Short action potential

Cardiac Muscle

- Myogenic

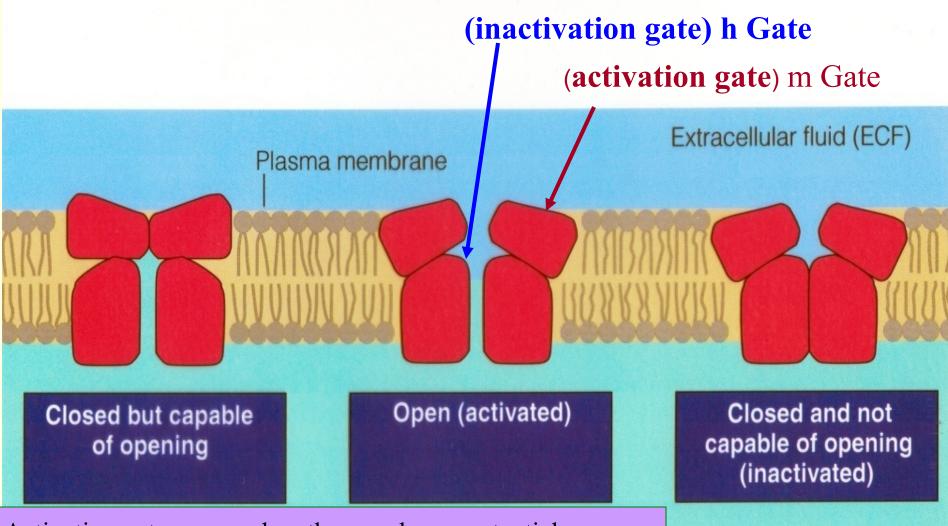
 (action potential originates within the muscle)
- Gap-junctions
- Action potential is longer

University of Jordan

Functional importance of Cardiac action potential

- The decrease in conductance (permeability) of potassium at phase 0 and 1 of the cardiac action potential contributes to the maintenance of depolarization in phase 2 (plateau)
- The long absolute refractory period prevent the occurrence of tetanus (maintained contraction without a period of relaxation) in the cardiac muscle.
- Skeletal muscle action potential is short that allows tetanus to occur

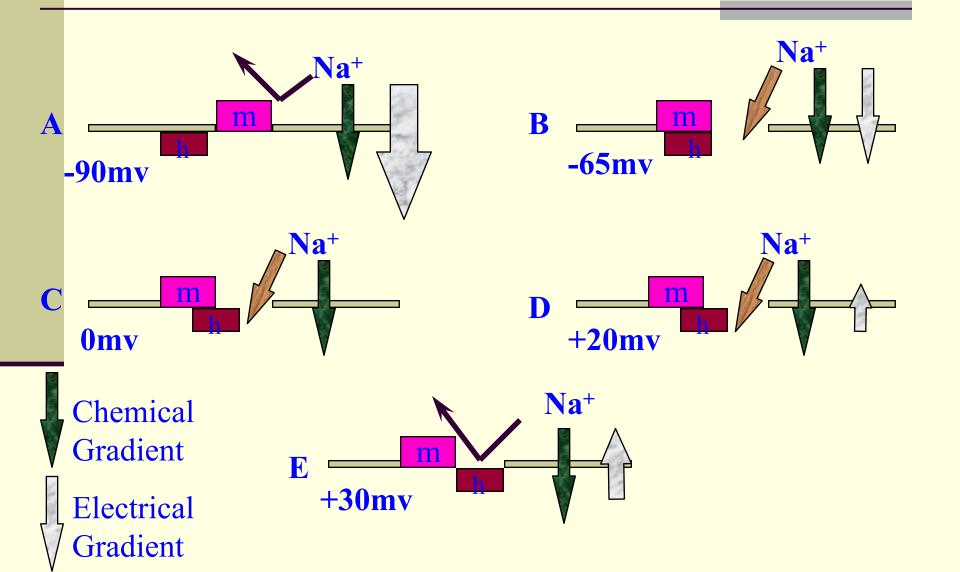
Conformations of a Voltage-Gated Na⁺ Channel

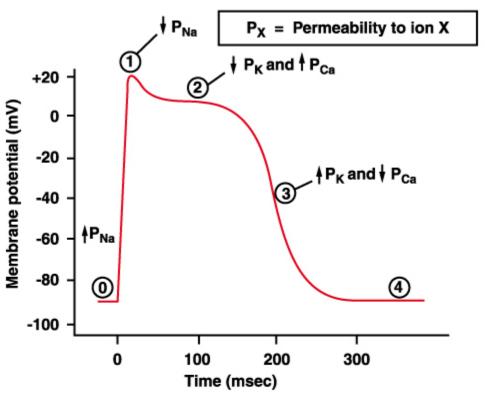


Activation gates open when the membrane potential becomes less negative and the inactivation gates close when the potential becomes less negative. The activation gate is fast but the inactivation is slow responding

Intracellular fluid (ICF)

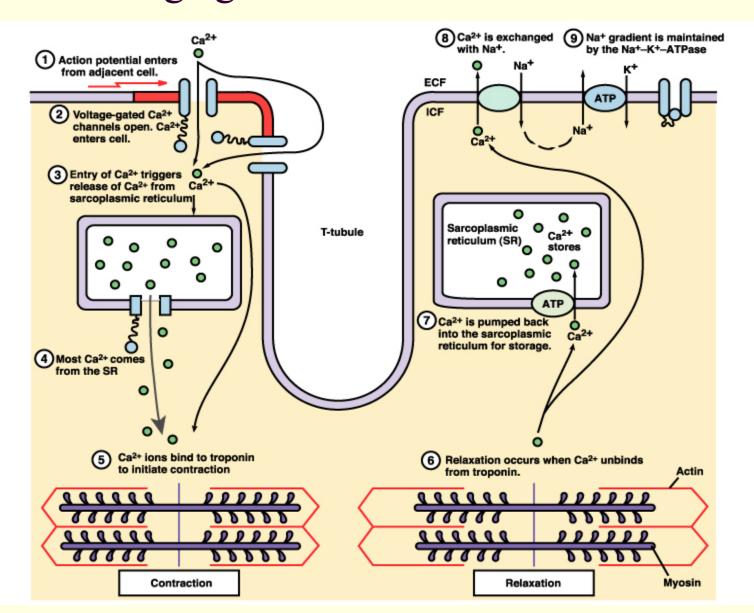
PHASE 0 OF THE FAST FIBER ACTION POTENTIAL



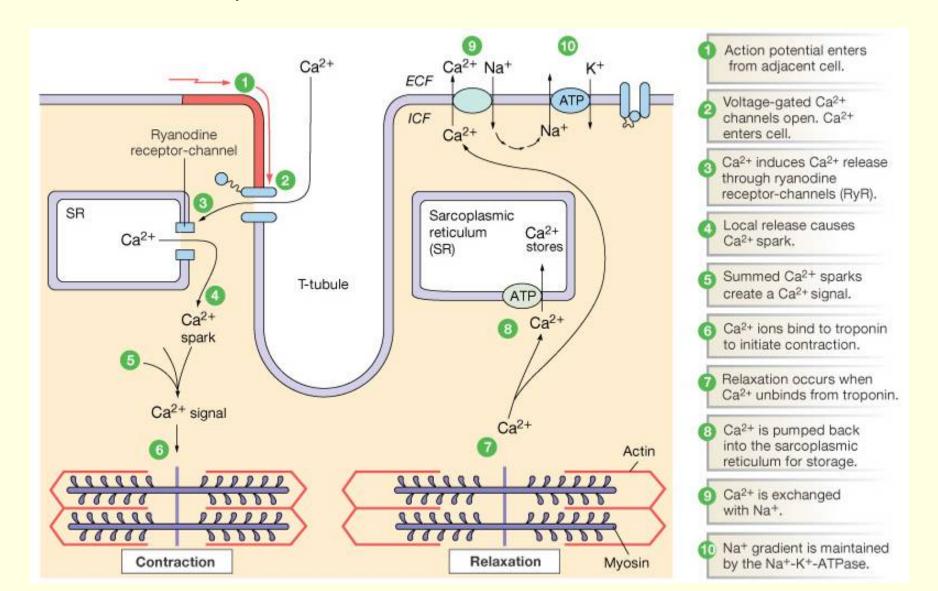


Phase	Membrane channels
0	Na+ channels open
1	Na+ channels close
2	Ca ²⁺ channels open; fast K ⁺ channels close
3	Ca ²⁺ channels close; slow K ⁺ channels open
4	Resting potential

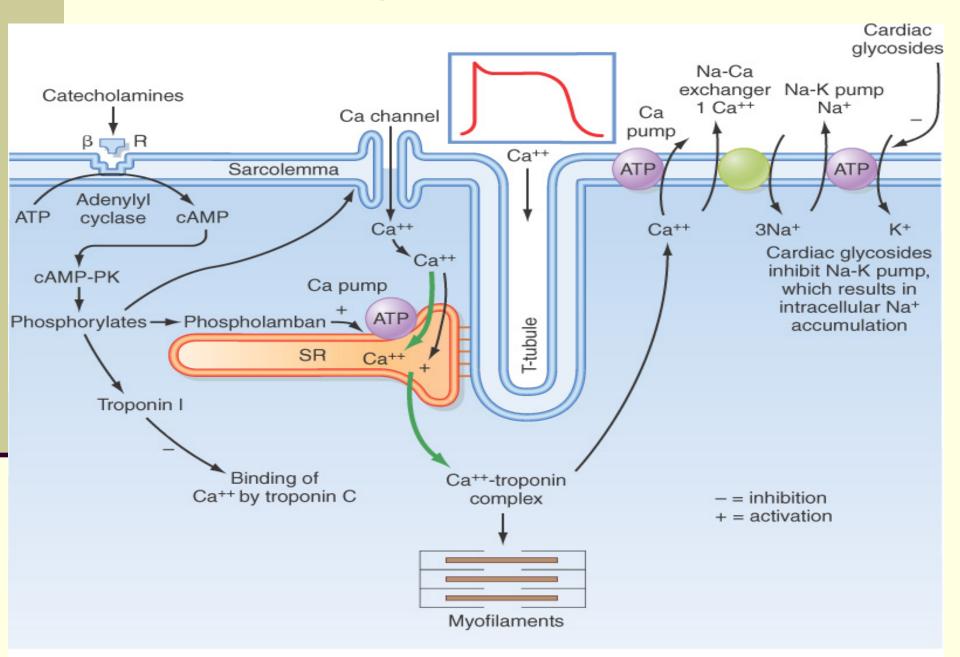
The importance of calcium influx through the slow voltage gated calcium channels



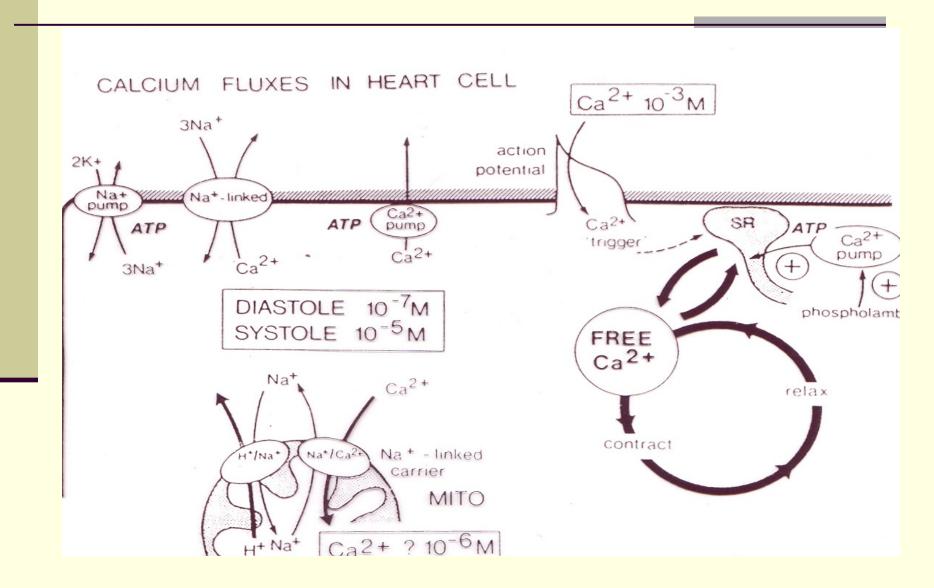
Mechanism of Cardiac Muscle Excitation, Contraction & Relaxation



Intracellular Calcium Homeostasis...1



Intracellular Calcium Homeostasis...2



Cardiac Muscle action potential Vs. Skeletal Muscle

- ➤ Phase 0 —Depolarization phase (Na⁺ influx)
- ➤ Phase 1 partial repolarization (Not in skeletal)
- ➤ Phase 2 Plateau (~ depolarization not in skeletal) slow calcium channels
- ➤ Phase 3 fast repolarization phase (K⁺ repolarization()
- > Phase 4 resting membrane potential

Thank You



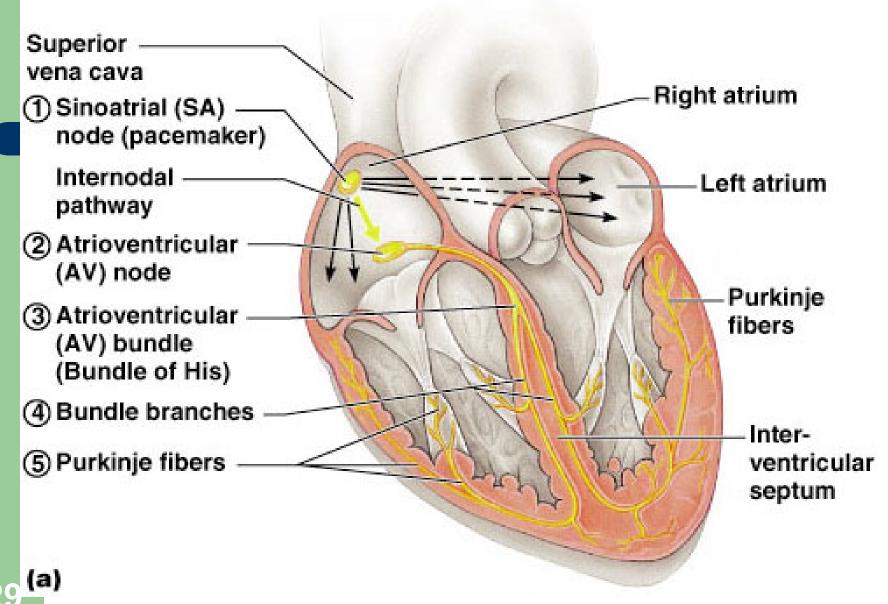
Conduction System of the Heart

Faisal I. Mohammed, MD, PhD

Objectives

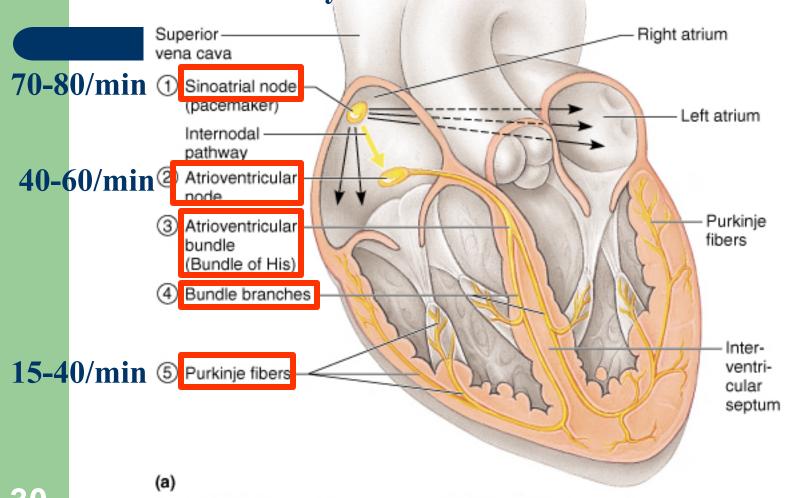
- List the parts that comprise the conduction system
- Explain the mechanism of slow response action potential (pacemaker potential)
- Point out the regulation of the conduction system potential by Autonomic Nerves

Conducting System of Heart



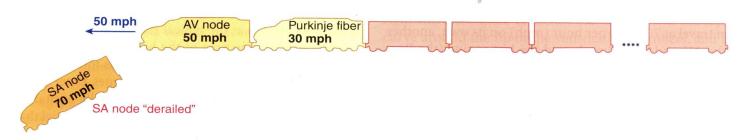
Intrinsic Cardiac Conduction System

Approximately 1% of cardiac muscle cells are autorhythmic rather than contractile

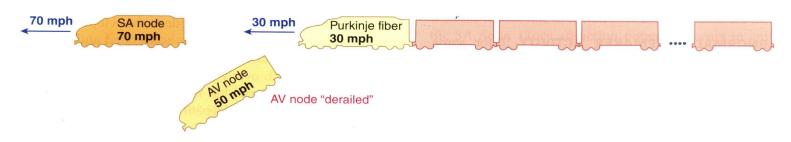




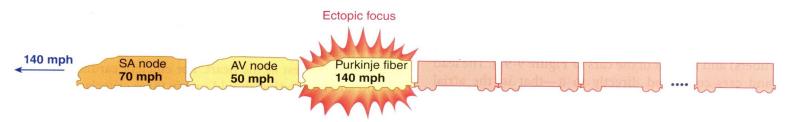
(a) Normal pacemaker activity: Whole train will go 70 mph (heart rate set by SA node, the fastest autorhythmic tissue).



(b) Takeover of pacemaker activity by AV node when the SA node is nonfunctional: Train will go 50 mph (the next fastest autorhythmic tissue, the AV node, will set the heart rate).



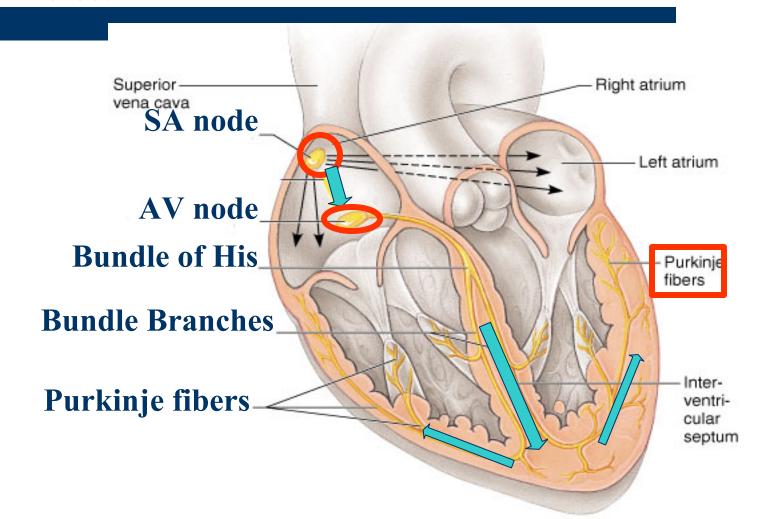
(c) Takeover of ventricular rate by the slower ventricular autorhythmic tissue in complete heart block: First part of train will go 70 mph; last part will go 30 mph (atria will be driven by SA node; ventricles will assume own, much slower rhythm).



(d) Takeover of pacemaker activity by an ectopic focus: Train will be driven by ectopic focus, which is now going faster than the SA node (the whole heart will be driven more rapidly by an abnormal pacemaker).

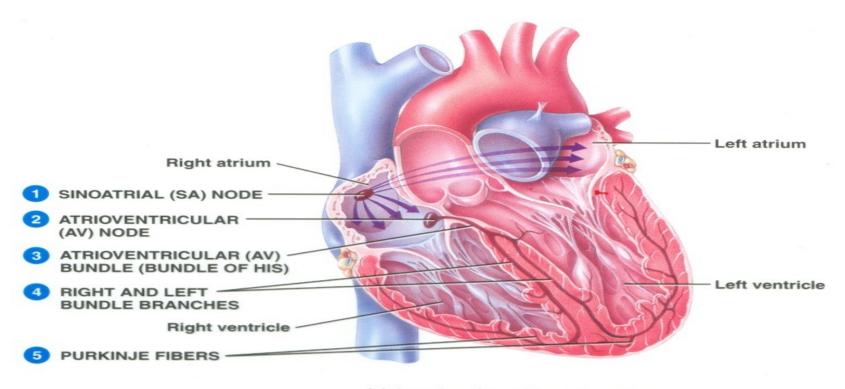
Intrinsic Conduction System

<u>Function</u>: initiate & distribute impulses so heart depolarizes & contracts in orderly manner from atria to ventricles.

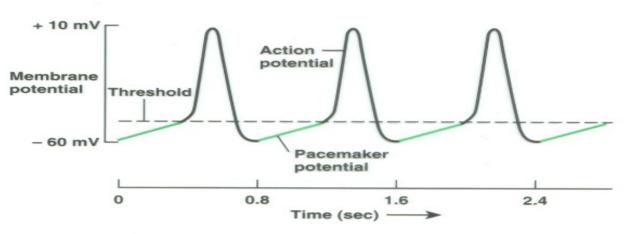


Sinus Node

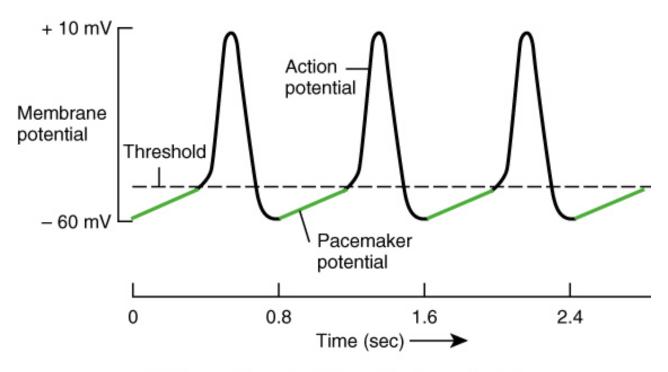
- Specialized cardiac muscle connected to atrial muscle.
- Acts as pacemaker because membrane leaks Na⁺ and membrane potential is -55 to -60mV
- When membrane potential reaches -40 mV, slow Ca⁺⁺ channels open causing action potential.
- After 100-150 msec Ca⁺⁺ channels close and K⁺channels open more thus returning membrane potential to -55mV.



(a) Anterior view of frontal section



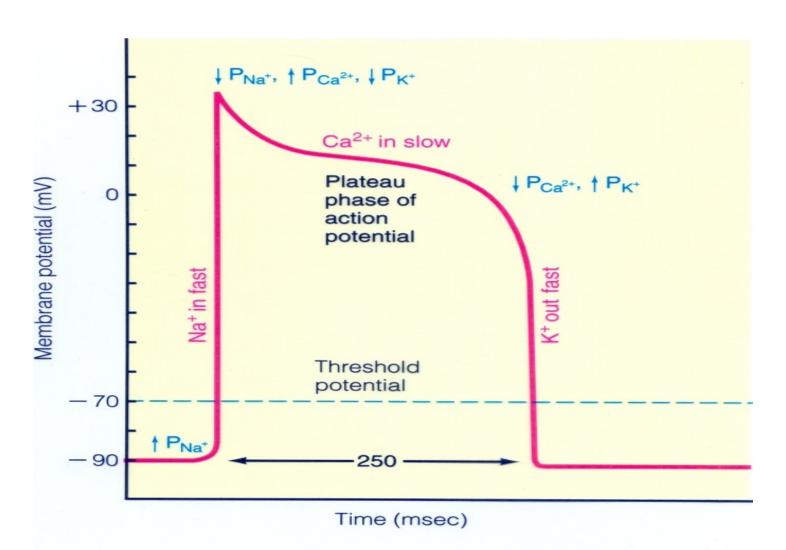
(b) Pacemaker potentials and action potentials in autorhythmic fibers of SA node



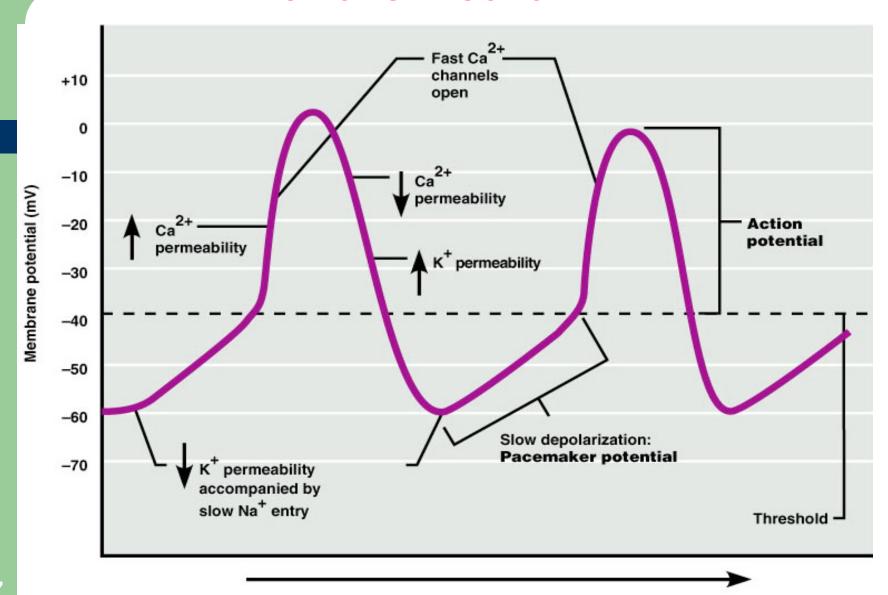
(b) Pacemaker potentials and action potentials in autorhythmic fibers of SA node

20.10b

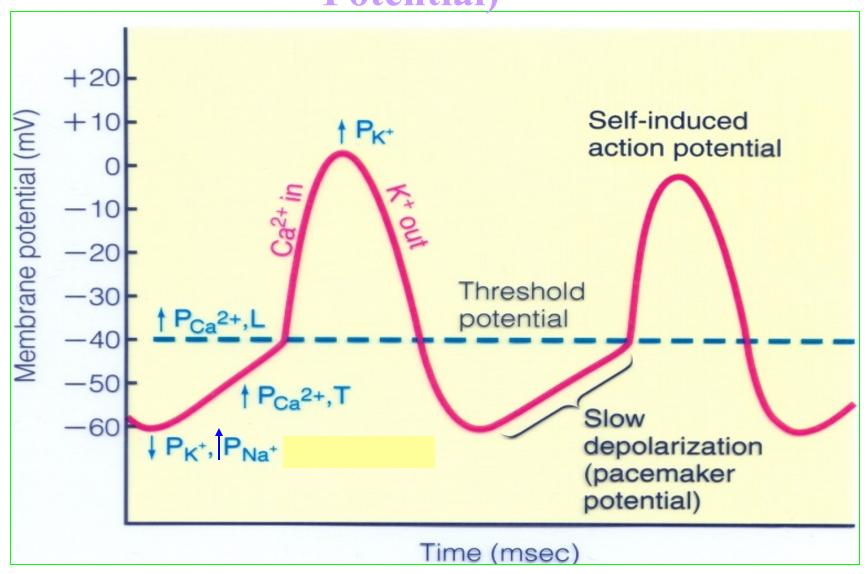
Fast Response Action Potential of Contractile Cardiac Muscle Cell



Pacemaker and Action Potentials of the Heart



Slow Response Action Potential (Pacemaker Potential)



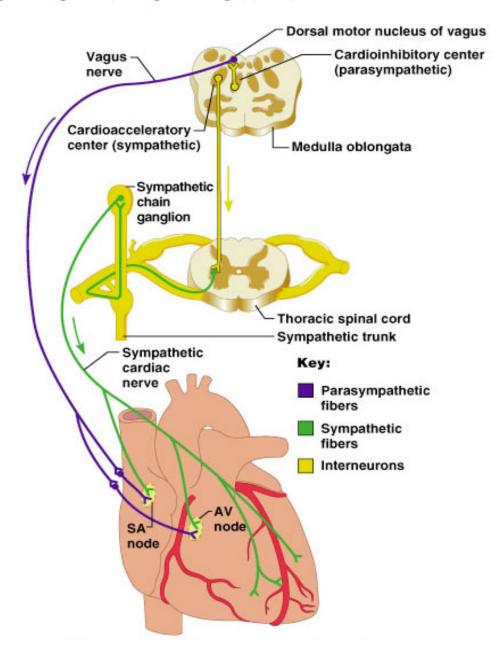
Intrinsic rate and speed of conduction of the components of the system

- SA node 60-80 action potential /min (*Pacemaker*)
- AV node 40-60 action potential /min
- Purkinje 15-40 action potential /min

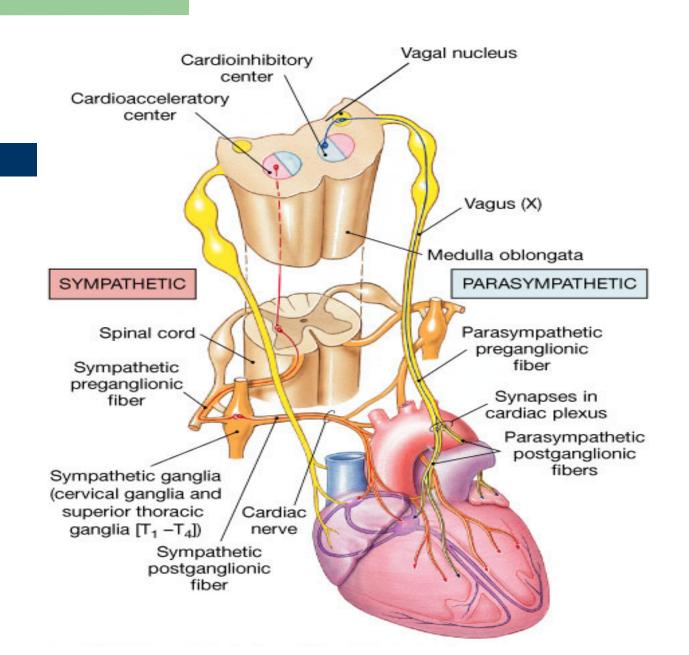
Conduction Speed

- SA node: slow speed of conduction
- Ventricular and Atrial muscle: Moderate speed
- AV node: slowest speed of conduction
- Purkinje fibers: Fastest speed of conduction
- Ectopic Pacemaker- Abnormal site of pacemaker

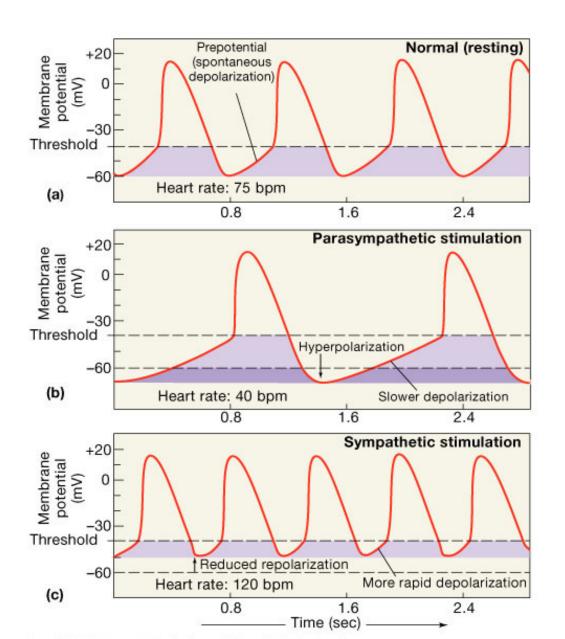
Extrinsic Innervation of the Heart



Autonomic Innervation of the Heart

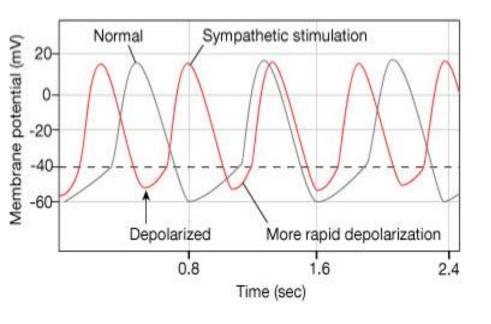


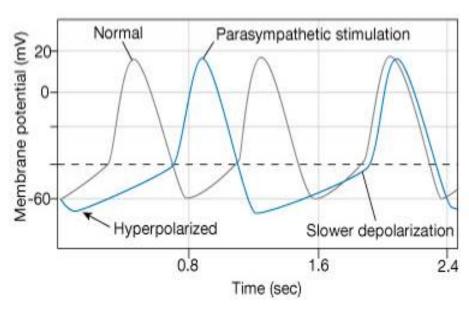
Pacemaker Function



Autonomic neurotransmitters affect ion flow to change rate

- **Sympathetic** increases heart rate by ↑ Ca⁺² & I_f channel (net Na⁺) flow
- **Parasympathetic** decreases rate by \uparrow K⁺ efflux & \downarrow Ca⁺² influx *What part of the graph is* <u>not</u> *changed by autonomic influences?*





Regulation of the heart beat

- Sympathetic from the cardiac plexus supplies all parts of the heart (atria, ventricle and all parts of the conduction system)
- Parasympathetic from Vagus nerves supply mainly the atria, SA and AV nodes, very little supply to ventricles
- Sympathetic: increase the permeability of the cardiac cells to Na⁺ and Ca⁺⁺ i.e Positive Chronotropic and positive Inotropic action
- Parasympathetic: Increase the permeability of the cardiac cells to K⁺ and decrease its permeability to Na⁺ and Ca⁺⁺

Sinus Node is Cardiac Pacemaker

- Normal rate of discharge in sinus node is 70-80/min.; A-V node 40-60/min.; Purkinje fibers 15-40/min.
- Sinus node is pacemaker because of its faster discharge rate
- Intrinsic rate of subsequent parts is suppressed by "Overdrive suppression"

Ectopic Pacemaker

- This is a portion of the heart with a more rapid discharge than the sinus node.
- Also occurs when transmission from sinus node to A-V node is blocked (A-V block).

Parasympathetic Effects on Heart Rate

- Parasympathetic (vagal) nerves, which release acetylcholine at their endings, innervate S-A node and A-V junctional fibers proximal to A-V node.
- Causes hyperpolarization because of increased K⁺ permeability in response to acetylcholine.
- This causes decreased transmission of impulses maybe temporarily stopping heart rate.

Sympathetic Effects on Heart Rate

- Releases norepinephrine at sympathetic ending
- Causes increased sinus node discharge (Chronotropic effect)
- Increases rate of conduction of impulse
 (Dromotropic effect)
- Increases force of contraction in atria and ventricles (*Inotropic effect*)

Thank You

