

# Screening Method of Anti-angina



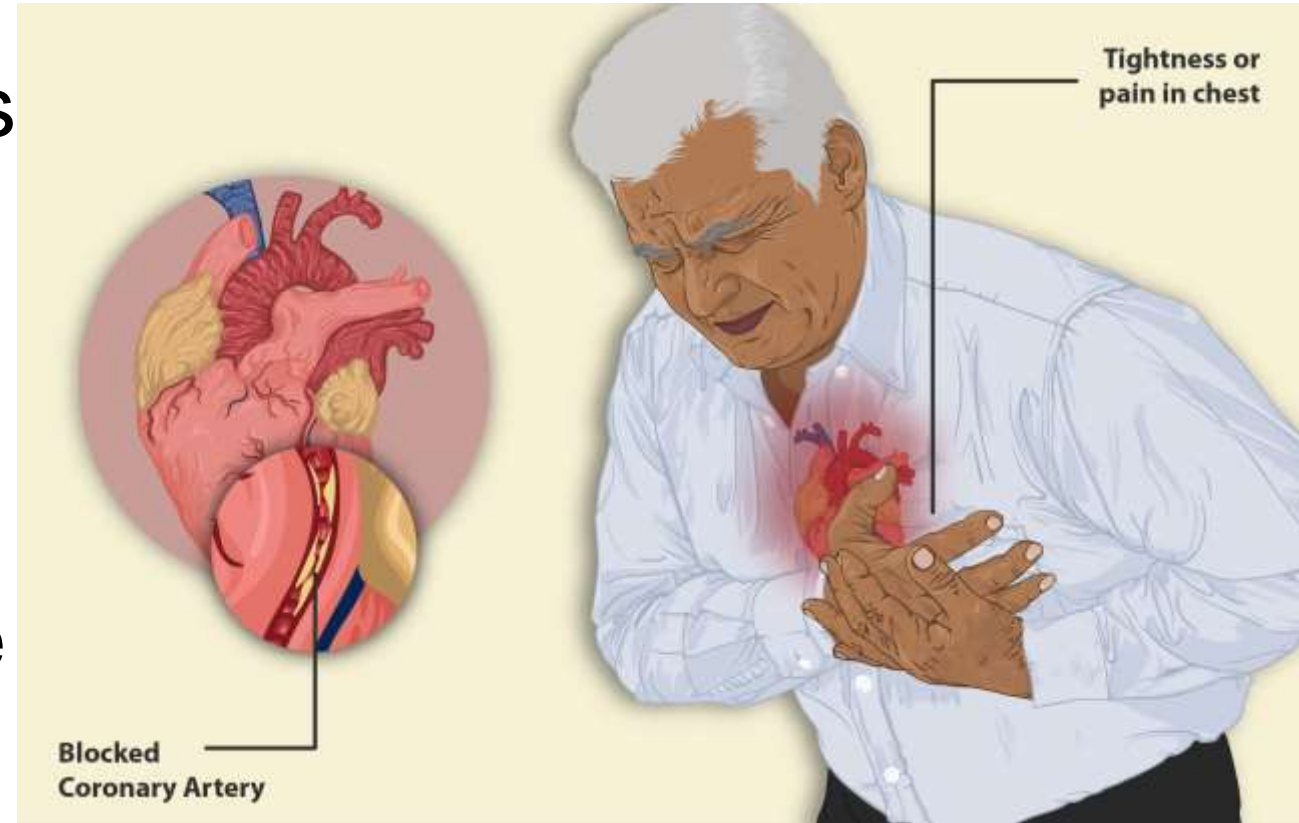
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# Introduction

- Angina pectoris is a symptoms of ischemic (less blood supply) heart disease.
- It is due to an imbalance between oxygen supply and oxygen demand of the myocardium.



# Cause of Angina

- Coronary Artery Disease
- Atherosclerosis
- Smoking
- High Blood Pressure
- High Cholesterol
- Diabetes
- Family History
- Obesity
- Lack of physical activity
- Stress



# Symptoms of Angina

- Weakness
- Heartburn
- Sweating
- Shortness of breath
- Anxiety
- Chest pain
- Indigestion



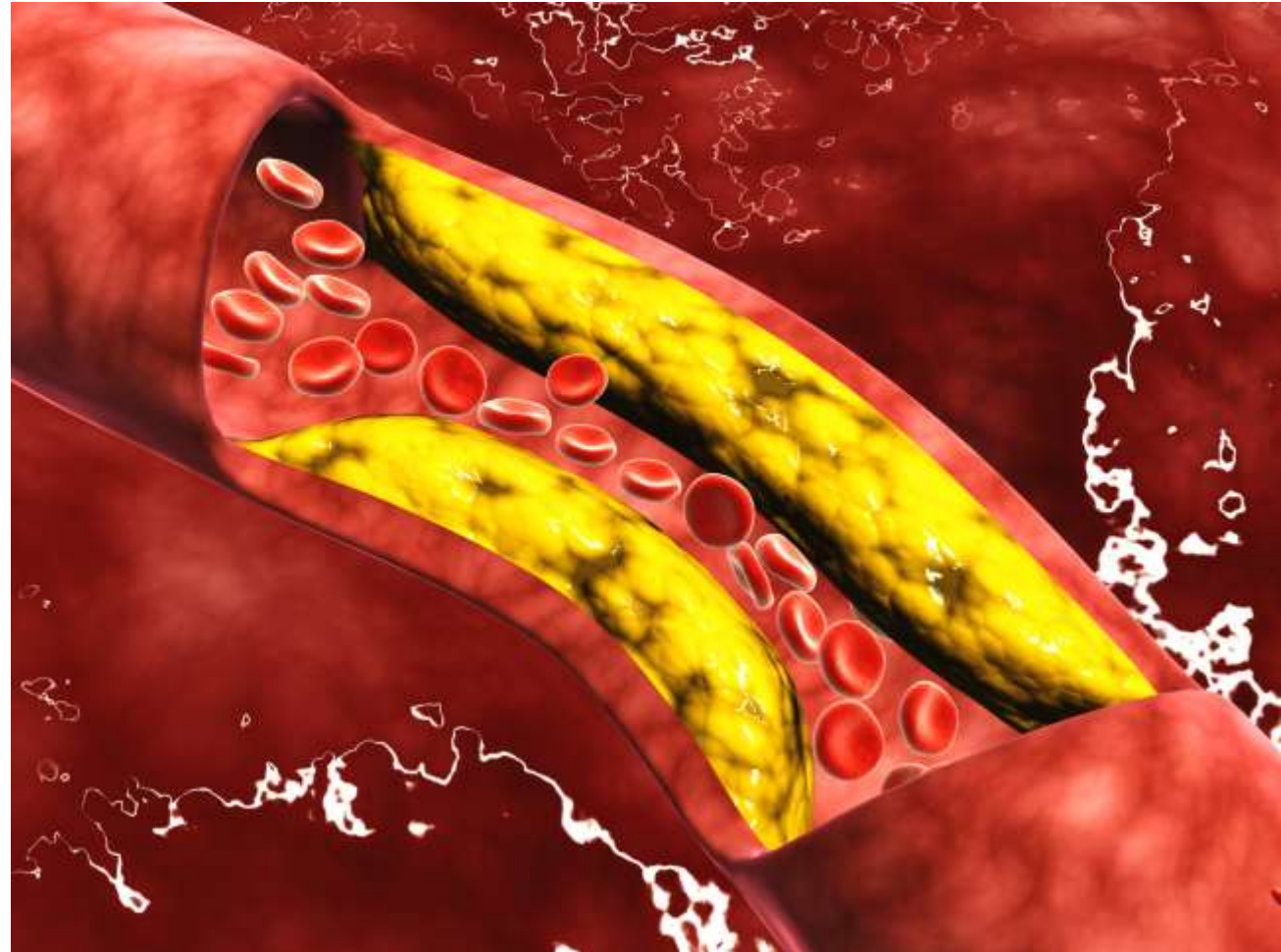
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# Types of Angina

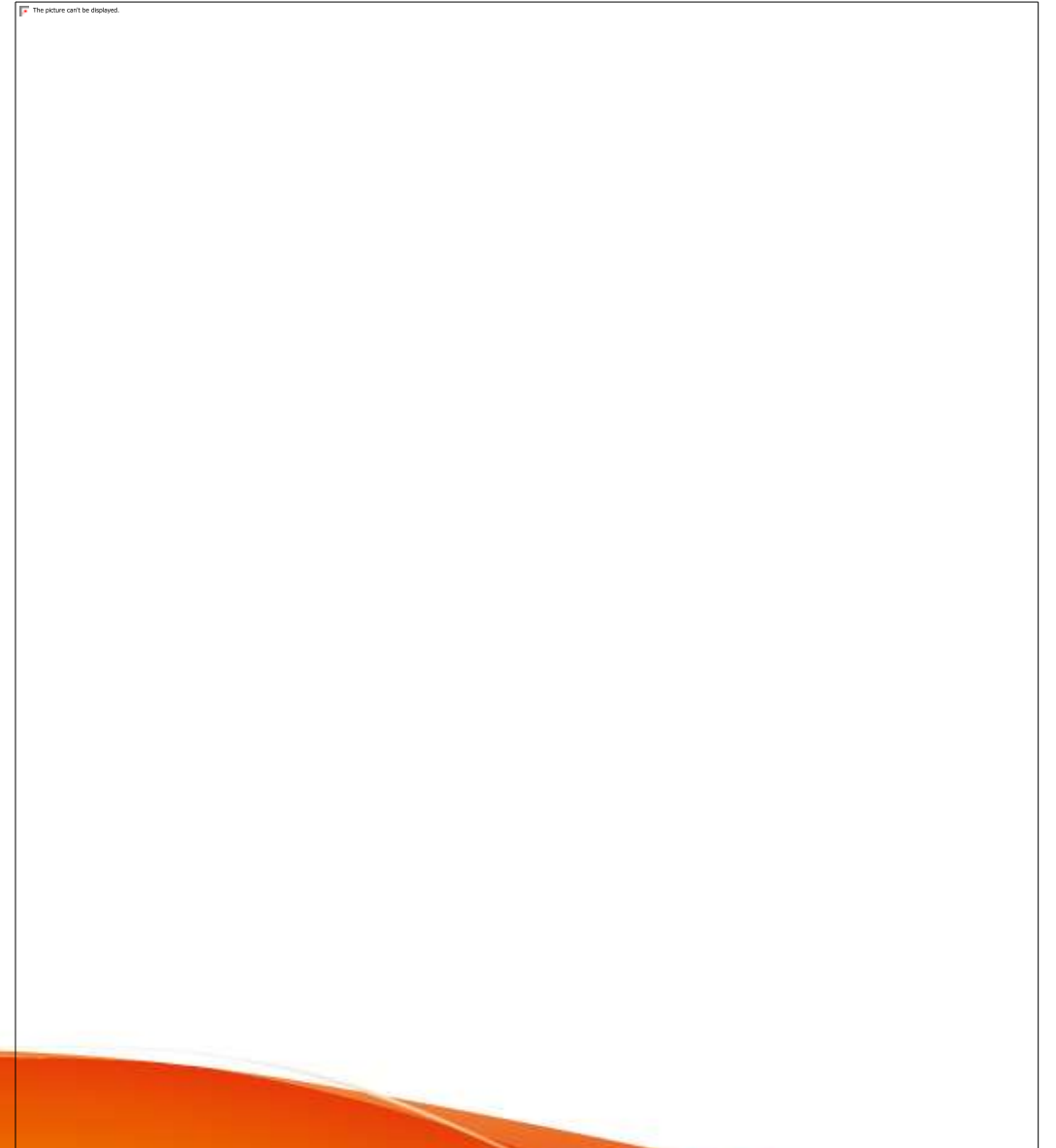
## Classic Angina(Angina of Exercise)-

- This angina pain occurs when the demand of oxygen exceeds the supply of oxygen most commonly due to diminished coronary flow.



# Vasospastic(Prinzmetal's or Variant) Angina

- This angina pain occurs at rest and is characterised by reversible coronary vasospasm, which in turn reduces the supply of oxygen.



# **Classification of Anti-angina Drug**

## **Nitrates-**

- **Long acting-** Isosorbide dinitrate(oral) Isosorbide mononitrate.
- **Short acting-** Glyceryl trinitrate, Isosorbide dinitrate(sublingual)

**B-blocker-** Propranolol, Metoprolol, Atenolol.

## **Calcium channel blocker-**

Verapamil, Amlodipine, Diltiazem.

**Pot.channel opener-** Nicorandil.

**Other anti-anginal-** Trimetazidine, Ranolazine, Aspirin.







# **PRECLINICAL EVALUATION**



# IN VITRO

- Isolated heart (Langendorff) technique
  - Isolated heart-lung preparation
  - Relaxation of bovine coronary artery
  - Coronary artery ligation in isolated rat heart
  - Isolated rabbit aorta preparation
  - Plastic costs technique in dogs
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# IN VIVO

- Coronary artery occlusion
  - Isoproterenol-induce myocardial necrosis
  - Stenosis-induce coronary thrombosis model
  - Electrical stimulation-induced coronary thrombosis
  - Myocardial ischemic preconditioning model
  - Electromagnetic flow meter
  - Occlusion of coronary artery
- 

# **IN VITRO**

## **Isolated heart(Langedorff) technique-**

**Principal-** is to maintain the heart perfused at constant temperature.

- Established in 1897 by Oscar Langendorff

**Requirments-** Guinea pigs (300-500gm)

Cold perfusion solution(4c)


Oxygentaed ringer solution

**Equipments-** Thoracic cage

Chronometer

Cannula, Double wall plexiglass perfusion apparatus, Small steel hook, Force transducer, polygraph.

# Procedure

- Guinea pigs of either sex weighing 300 to 500gm used for study and they are sacrificed by stunning.
  - Diaphragm is assessed by transabdominal incision and cut carefully to expose, the thoracic cavity.
  - Thoracic is opened by bilateral incision along the lower margins of last to first ribs.
  - The heart is cradled between finger and lifted before incising the aorta, enclave pulmonary veins.
- 



- Immediately after excision, heart is dipped in cold perfusion solution .The aorta is located and cut below the point of division.
- Cannula is inserted into the aorta and tied the heart perfused with oxygenated ringer's solution.
- The heart is transferred to a double wall plexiglass perfusion apparatus maintained at 37c,40mm Hg pressure.
- Small steel hook with a string is attached to apex of the heart.
- Contractile force is measured isometrically by a force transducer and recorded on a polygraph.

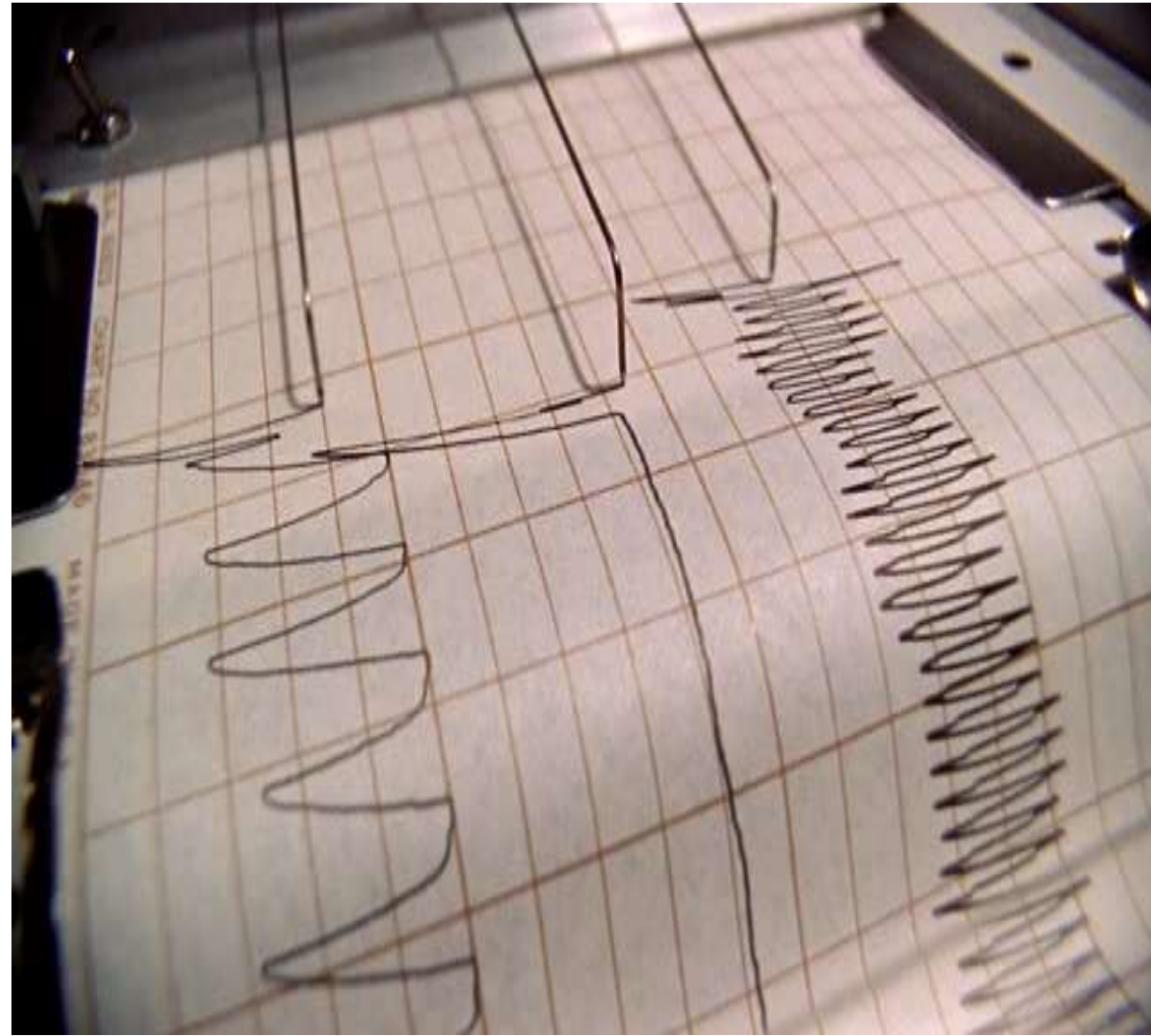
- Heart rate is measured through a chronometer coupled to the polygraph.
- Then drugs(standard,test) are injected into the perfusion medium.
- The atherosclerosis effect of test drug is indicated by an increase in coronary blood flow.

**Conclusion-** The incidence and duration of ventricular fibrillation coronary flow in atrophic state and K<sup>+</sup> levels after treatment with drug are compared with control.

- Heart rate by chronometer-attached to polygraph.
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- Contractile force is measured isometrically by a force transducer.
- Cardiac output determined by electromagnetic flow probes in the outflow system.



# **Isolated heart-lung preparation**

**Aim-** The isolated heart lung preparation is used to study various physiological and pharmacological processes.

**Animal required-** wistar rat (300-500gm)


**Chemical required-** Pentobarbitone sodium

Ice-cold saline

Krebs ringer, bicarbonate buffer

**Equipments required-** Cannula, Artificial respiration, Electrical amplifier.

# Procedure

- Wistar rat is anaesthetized with pentobarbitone sodium(50mg/kg).
  - The Trachea is cannulated animals is maintained on artificial respiration.
  - The chest cavity is opened and ice - cold saline is injected to arrest the heart.
  - The aorta, superior and inferior vena cava are cannulated.
  - The heart lung preparation is perfused with Krebs-Ringer bicarbonate buffer (pH-7.4) containing rat RBC (hematocrit 25%).
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- The perforate is pumped from the aorta and is passed through the pneumatic resistance and collected in a reservoir maintained at 37c.
- It is then returned to inferior vena cave thus perfusing only the heart and the lung.
- Test drug is administered into the perforate 5min after start of experiment.



- Cardiac output is recorded with an electromagnetic blood flow meter and mean arterial pressure from the pneumatic resistance.
- with the help of a bio electrical heart rate is recorded.

**Conclusion-** Hemodynamic data and recovery time of the test drug group and control group(without any treatment) is compared using ANOVA and Kruskal-Wallis test respectively.