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#### Coagulation cascade and hemostatis

Bhagyalakshmi K

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#### Coagulation Cascade and Hemostasis

Dr. Bhagyalakshmi K Professor & Head Department of Physiology KMC - Mangalore

## **Objectives**

- 1. List the coagulation factors
- 2. Describe the steps involved in the mechanism of clot formation
- 3. Role of platelets in hemostasis

#### Hemostasis

Hemostasis is the arrest of bleeding

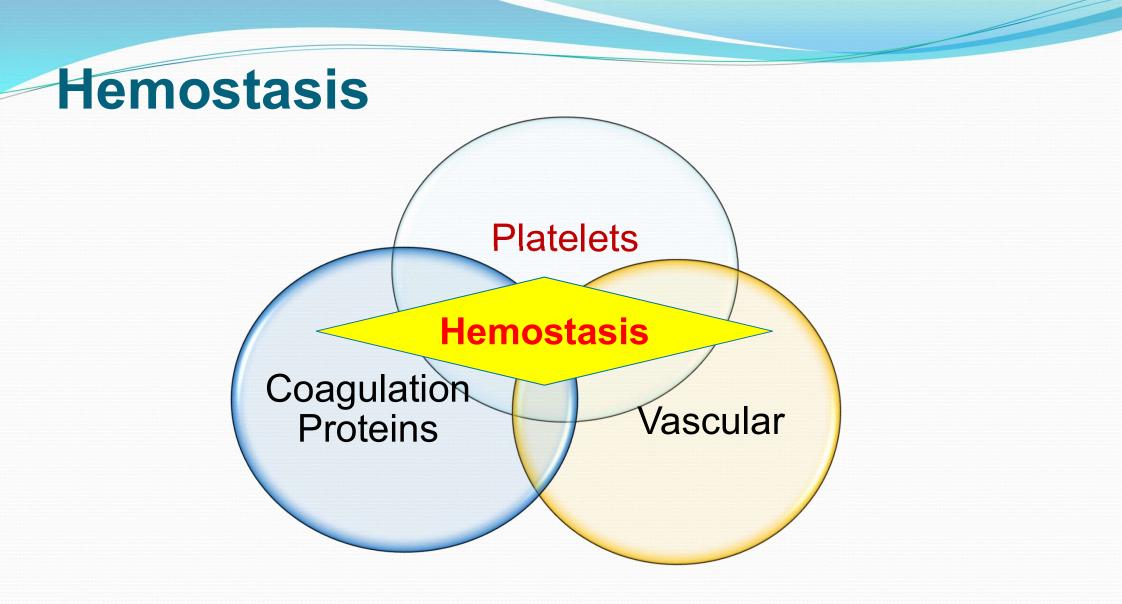
 Hemostasis: Process of forming clots in walls of damaged blood vessels and preventing blood loss while maintaining blood in a fluid state within the vascular system

#### Hemostasis

Physiological balance exists between

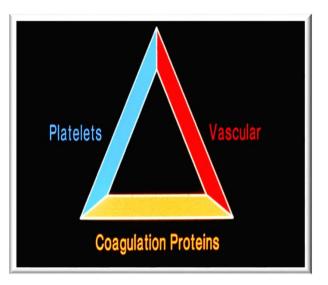
- Factors promoting coagulation (Procoagulants)
- Factors inhibiting coagulation (Anticoagulants)





#### Hemostasis

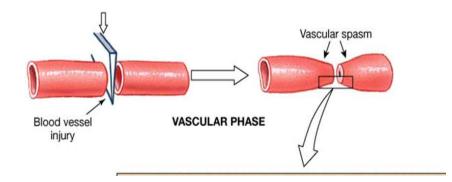
- 1.Vasoconstriction
- 2.Platelet plug formation
- 3.Coagulation of blood



## Vasoconstriction

- Constriction of blood vessel- local phenomenon,
- Nerve plexus
- Local myogenic spasm
- Local chemical factors
  - Serotonin
  - Thrombaxan A2 released from platelets

Collagen fibers exposed in damaged blood vessel favors platelet plug formation



#### **Properties of Platelets**

When a blood vessel is injured,

1. Adhesion: Platelets adhere to exposed collagen&Von Willebrand factor

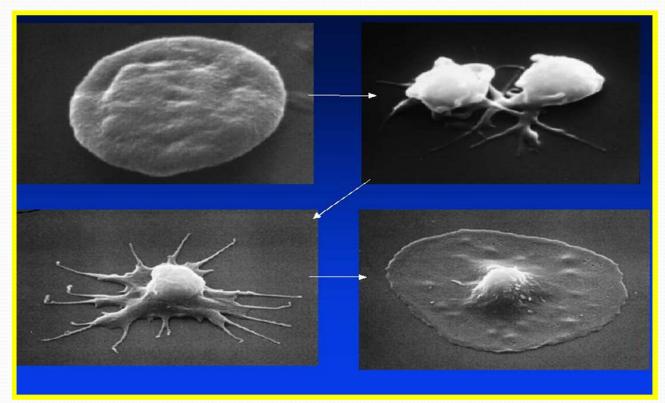
#### 2.Activation:

- Platelets binding to collagen -platelet activation
- Activated platelets swell up, sticky, loose their shape, send pseudopodia
- Degranulate & attract more platelets

#### 3. Aggregation:

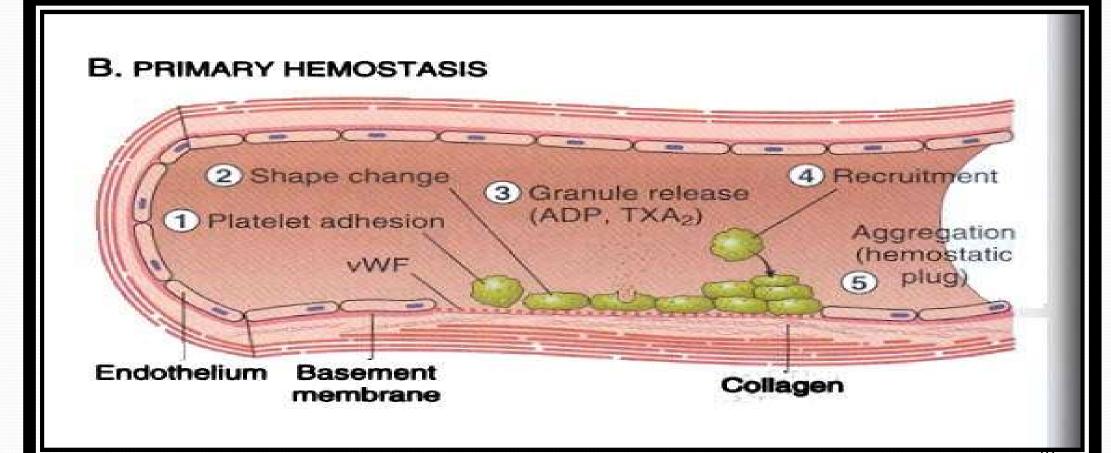
 Released ADP, Platelet- activating factors act on other platelet to produce further accumulation

## **Properties of Platelets**



#### Platelets change their shape

#### **Primary Hemostasis**



## **Platelet plug formation**

Platelets aggregate and form a temporary loose plug to prevent blood loss--Platelet Plug- Primary hemostasis



#### **Blood coagulation or clotting**

- Process of conversion of fluid blood into semisolid jelly like mass is known as coagulation or clotting of blood
- Occurs over temporary hemostatic plug
- Must for permanent arrest of bleeding

#### **Coagulation of blood: Clotting factors**

- I- Fibrinogen,
- II- Prothrombin,
- III- Tissue thromboplastin,
- IV- Calcium,

- VIII- Anti haemophilic factor, AHF-A
- IX- Christmas factor, AHF-B
- X- Stuart prower factor,
- XI- Plasma Thromboplastin Antecedent, AHF-C
- V- Labile factor, Proaccelerin, XII- Hagemans factor, Glass factor No factor VI
  - VII- Stable factor Proconvertin XIII- Fibrin stabilizing F, Laki-Lorand F

Foolish People Try Climbing Long Slopes, After Christmas Some People Have Fallen

## **Coagulation of blood**

#### **Clotting factors:**

VIII- Anti haemophilic factor, AHF-A

- IX- Christmas factor, AHF-B
- X- Stuart prower factor,
- XI- Plasma thromboplastin antecedent, AHF-C
- XII- hagemans factor, Glass factor
- XIII- fibrin stabilizing F, Laki-Lorand F

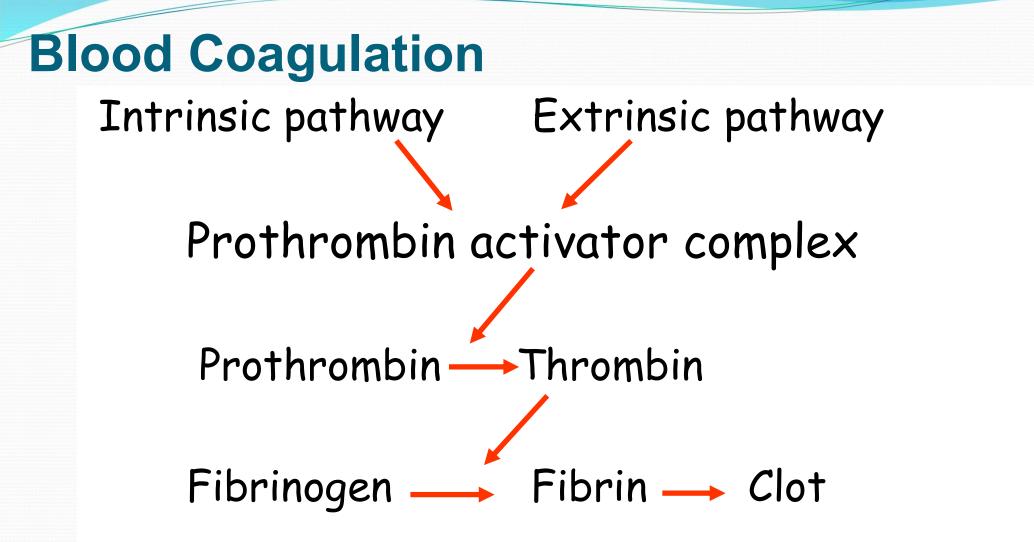
#### **Clotting Cascade**

- Factors, act like a row of dominoes toppling against each other
  Create a chain reaction
- If one factor is missing, chain reaction cannot proceed



## **Steps of coagulation**

- 1. Prothrombin activator complex formation
- 2. Conversion of prothrombin to thrombin
- 3. Conversion of fibrinogen to fibrin & clot formation



## Question

 Explain the mechanism of blood coagulation, when blood taken in a glass tube.

#### **Blood Coagulation**

Intrinsic pathway: When blood exposed to

- 1. Damaged endothelial cells
- 2. Collagen fibers underlying endothelium
- 3. Trauma to the blood itself
- 4. Electro negatively charged wettable surfaces like glass

#### **Blood Coagulation**

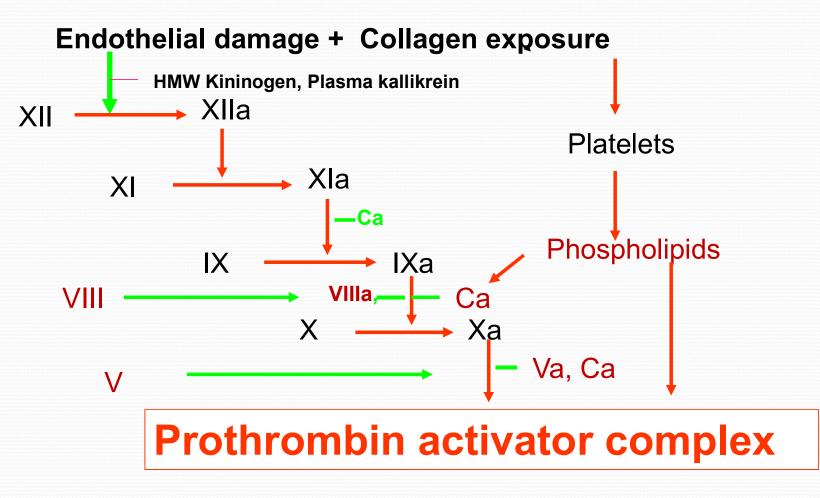
#### Extrinsic pathway:

- Begins with tissue thromboplastin (F-III)
- Released from the traumatized tissue

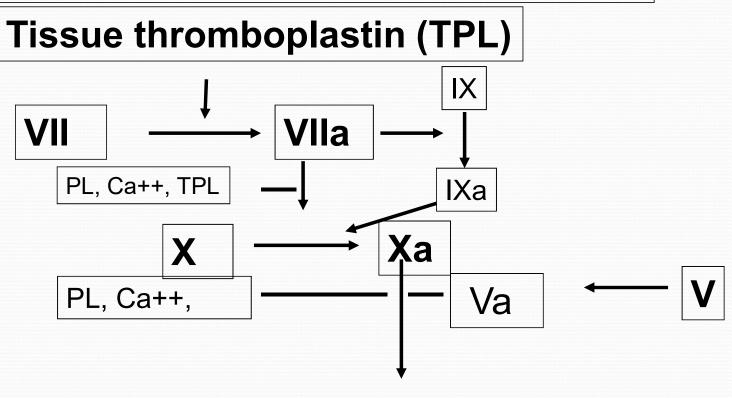
#### Clotting inside the body,

- Involves both intrinsic & extrinsic systems
- they both complement each other

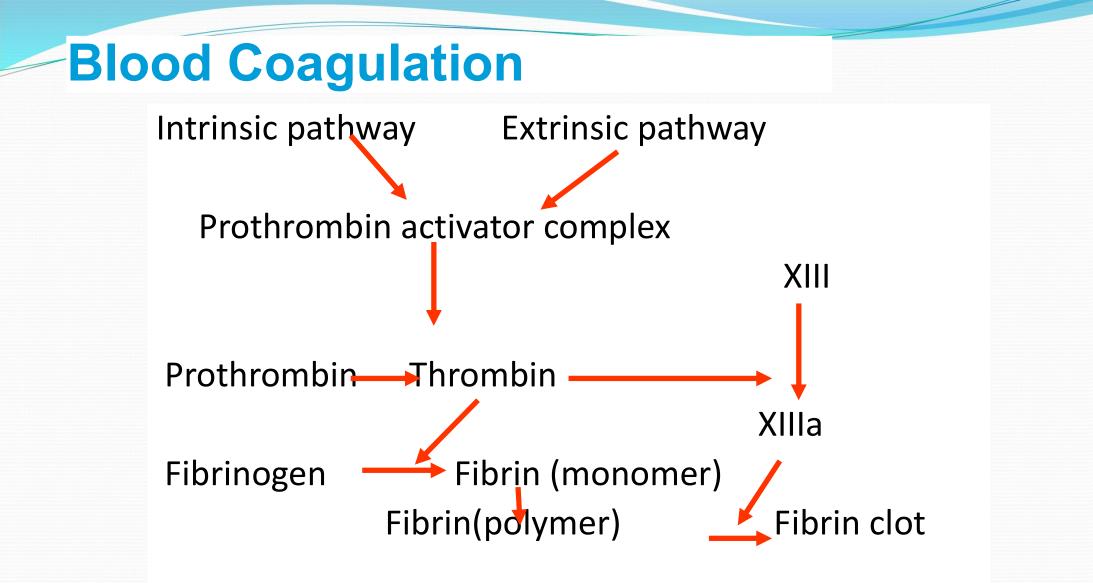
#### Intrinsic pathway



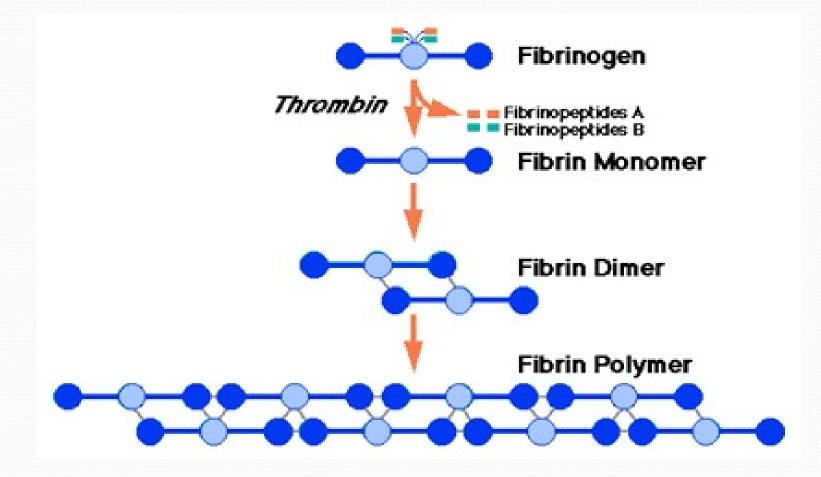
## **Extrinsic Pathway**



#### **Prothrombin activator complex**



#### **Blood Coagulation**

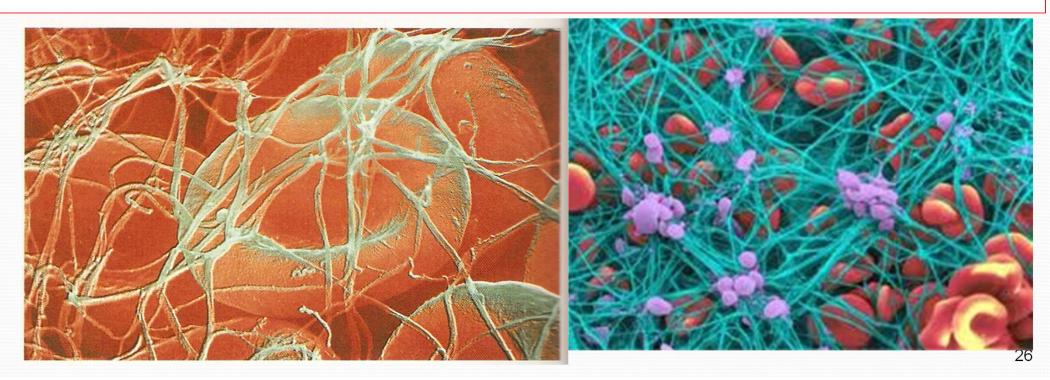


#### **Clot retraction**

- Fibrin threads, attaches to platelet plug & damaged vessels
- Contraction of platelet spicules attached to fibrin clot retraction
- As clot retracts edges of cut blood vessels are brought together
- Help for ultimate state Hemostasis
- Clot retraction-20-60min
- 40- 60% decrease in size of clot

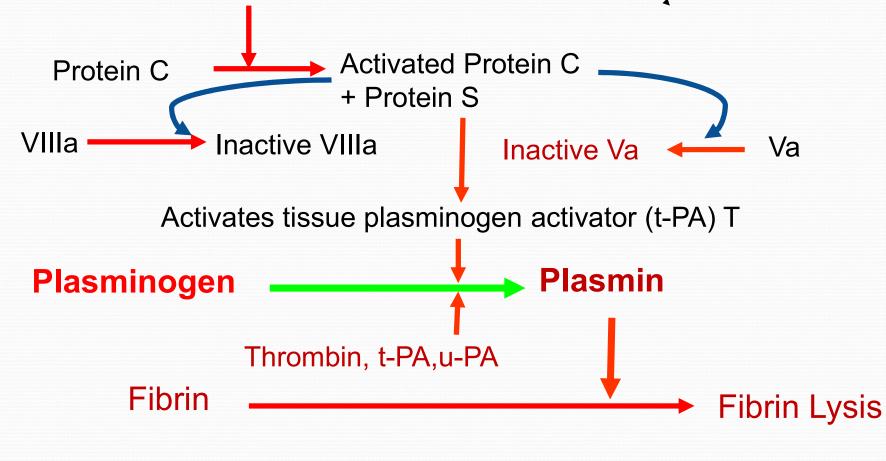
#### **Blood Clot**

- Meshwork of fibrin threads & entrapping blood cells and plasma
- Serum oozes out of it



#### **Fibrinolytic System**

#### **Thrombomodulin + Thrombin**



#### **Functions of Platelets**

- Role in Vasoconstriction
- Primary Hemostasis-by temporary Platelet Plug
- Secondary hemostasis-Help in clot formation
- Clot retraction: Due to actin, & Myosin
- Repair of capillary endothelium: PDGF- Multiplication of endothelial cells & fibroblasts

#### **Special attributes of Coagulation Process**

- Enzyme substrate reaction
- System acts as a bio amplifier system
- Cascade reaction
- There are +ve & –ve feedback mechanisms
- Ca++ are essential in almost all steps

## **Applied Physiology**

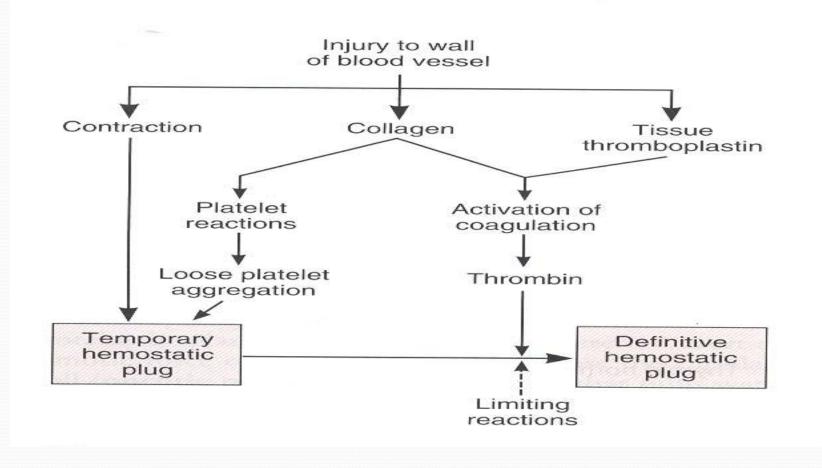
- Purpura: Immune thrombocytopenic Purpura
- Hemophilia
- Disseminated intravascular coagulation (DIC)

## **Applied Physiology**

**Commonly Performed Tests** 

- Bleeding time
- Clotting Time
- Prothrombin time
- Activated partial thromboplastin time (aPTT or APTT)
- Assay of individual clotting factors

# Summary -Hemostasis







# Thank you



- Bleeding time is prolonged in -----Thrombocytopenia
- Classic hemophilia is due to ----- Factor 8 deficiency
- Prothrombin time is the test for..... Extrinsic pathway
- Anticoagulant potency of Heparin increases by combining with .....antithrombin
- Fibrin stabilizing factor is mainly released from ....Platelets