

Blood Coagulation - Basics



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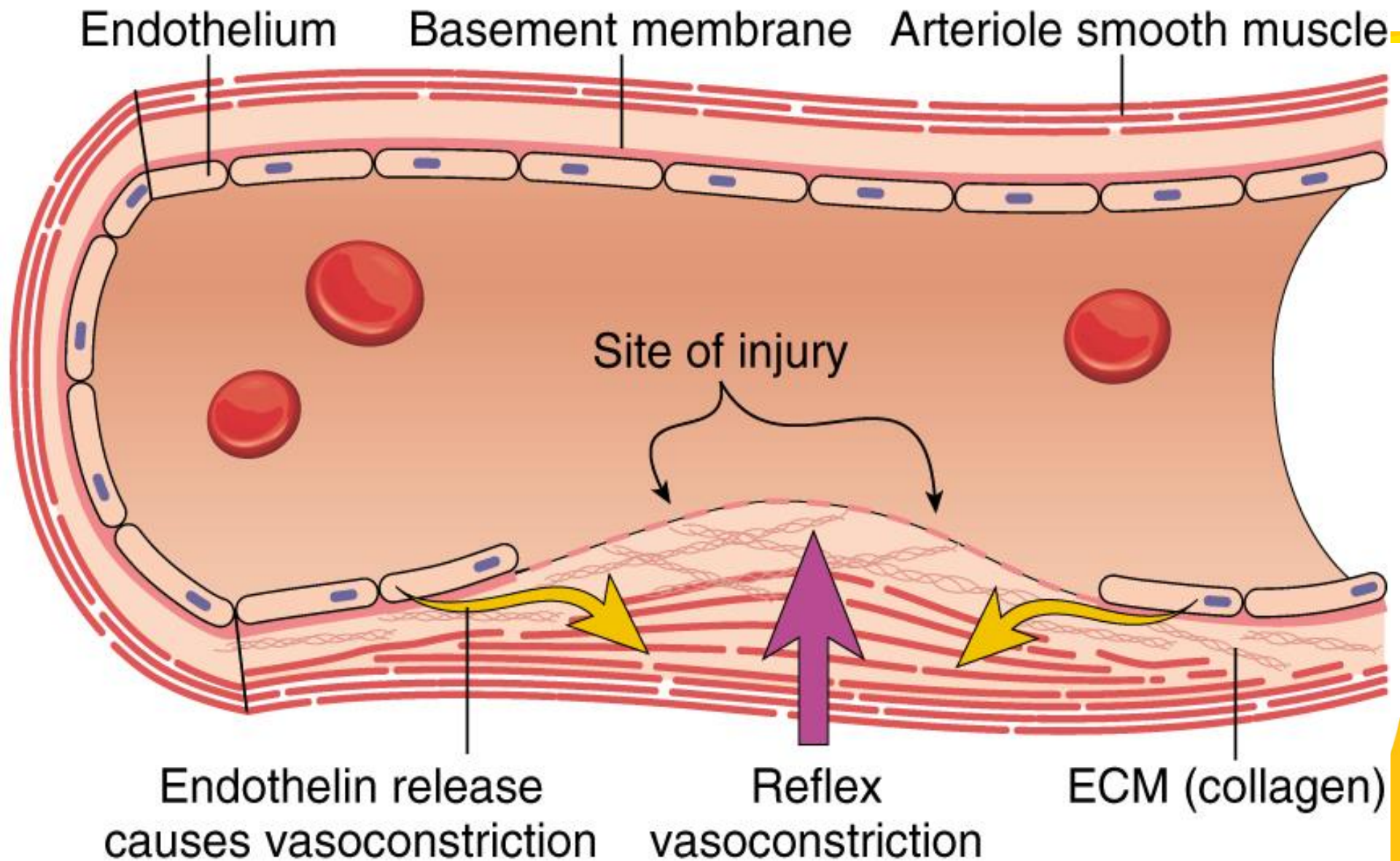
Basic Coagulation Profile

- Bleeding Time
- Platelet Count
- **Prothrombin Time (PT)**
- **Activated partial Thromboplastin Time (APTT)**
- Thrombin Time
- Fibrinogen Assays

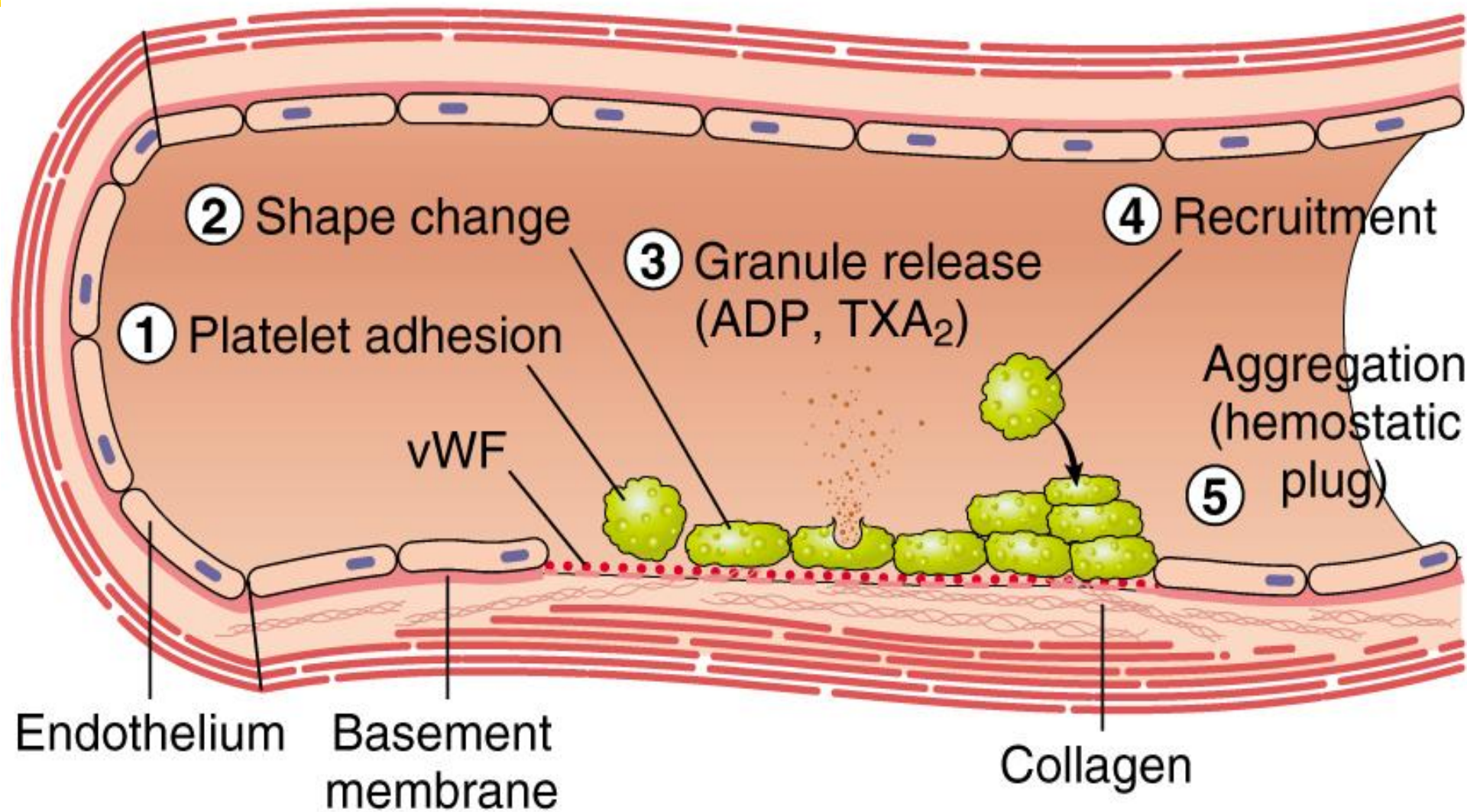
Hemostasis

- Hemostasis (“hemo”=blood; stasis= remain”) is the stoppage of bleeding, which is vitally important when blood vessels are damaged.
- Three steps:
 - Vasoconstriction
 - Primary Hemostasis- Platelet Plug formation
 - Secondary Hemostasis- Coagulation cascade activation leading to stable Clot formation

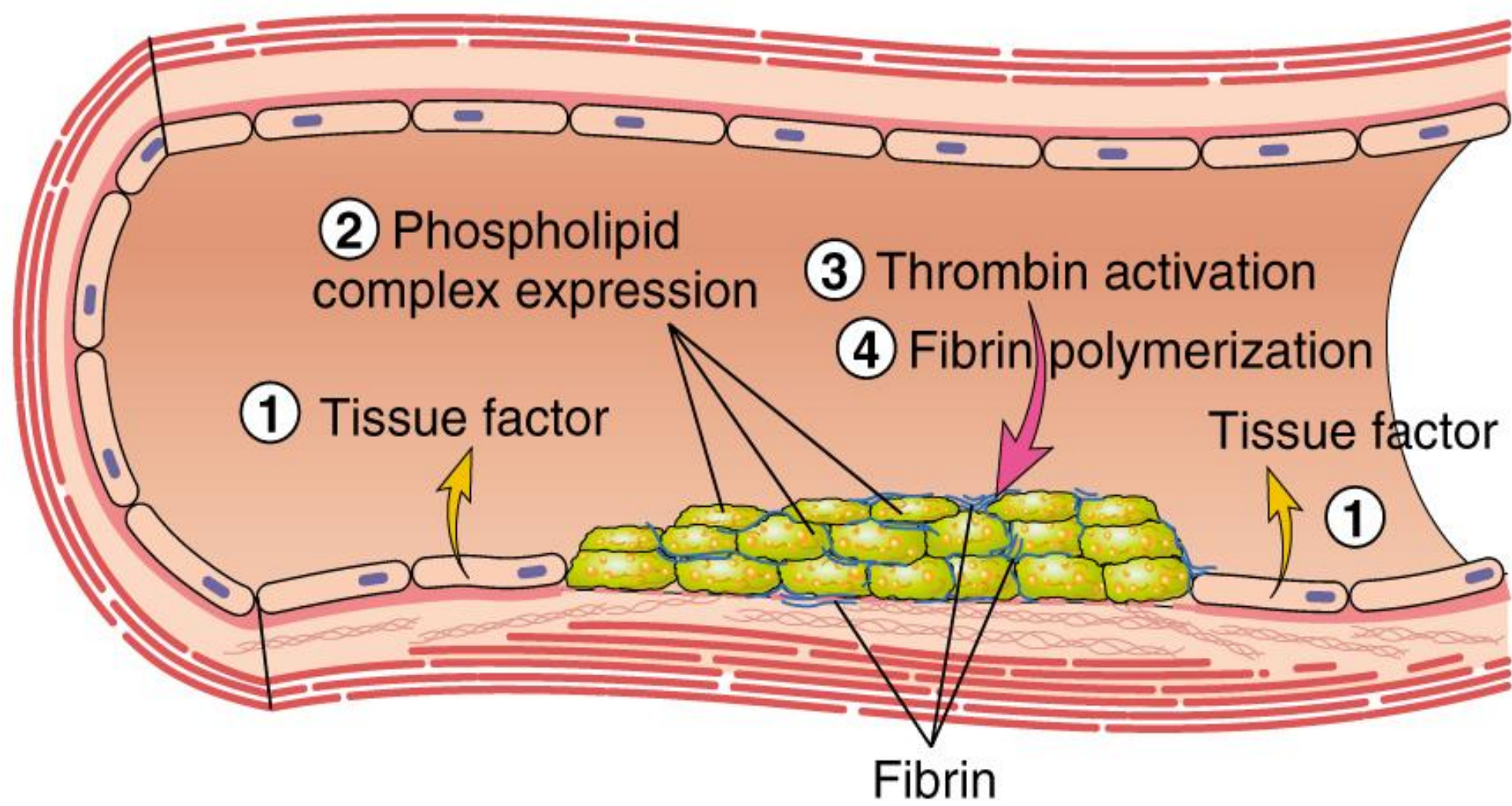
A. VASOCONSTRICTION



B. PRIMARY HEMOSTASIS

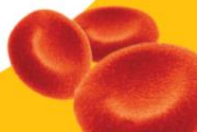


C. SECONDARY HEMOSTASIS



CLOTTING FACTORS

- I (fibrinogen)
- **II (prothrombin)**
- Tissue factor
- Calcium (Factor IV)
- V (proaccelerin, labile factor)
- VI
- VII (stable factor)
- **VIII (antihemophilic factor)**
- IX (Christmas factor)
- X (Stuart-Prower factor)
- XI (plasma thromboplastin antecedent)
- XII (Hageman factor)
- XIII (fibrin-stabilizing factor)
- von Willebrand factor



Coagulation made easy

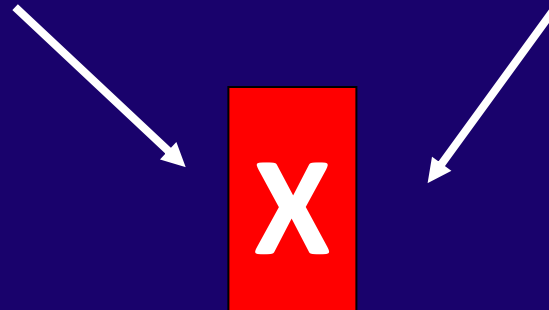
Intrinsic Pathway

Extrinsic Pathway

Coagulation made easy

The PTT Pathway
Intrinsic

The PT Pathway
Extrinsic

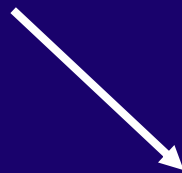


The PT and the PTT pathway meet at factor X, because "X" marks common pathway

Coagulation made easy

The PTT Pathway
Intrinsic

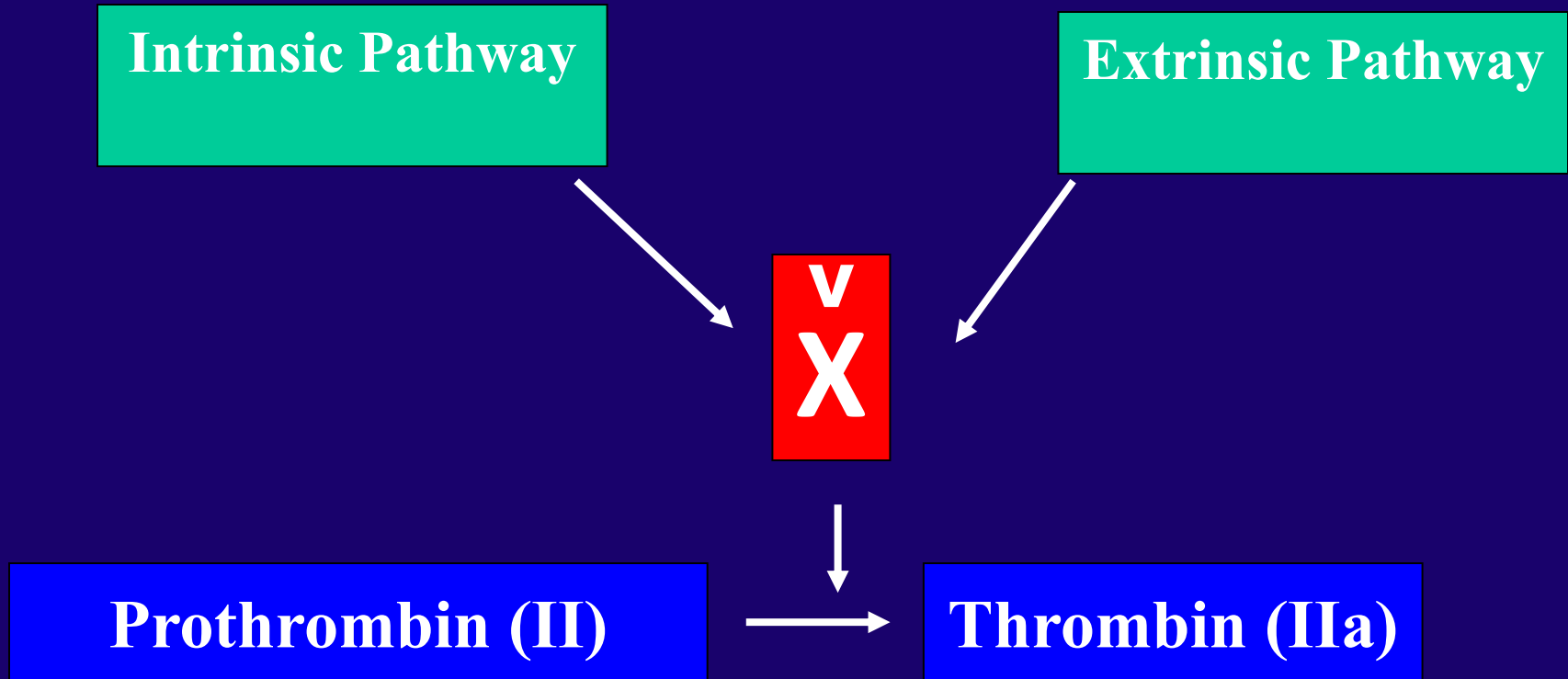
The PT Pathway
Extrinsic



Ca⁺⁺ , Phospholipid



Coagulation made easy



Coagulation made easy

Intrinsic Pathway

Extrinsic Pathway



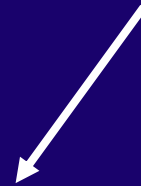
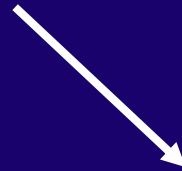
Thrombin, among other things, converts the soluble molecule fibrinogen into a solid fibrin clot

Prothrombin

Thrombin

Fibrinogen

Fibrin

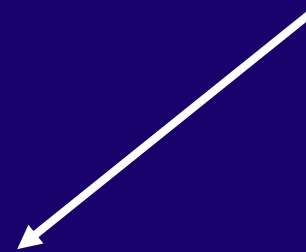
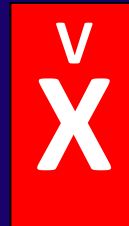


Coagulation made easy - the PT

The PT Pathway

WEPT 7

Warfarin
Extrinsic Pathway
PT
Factor VII



Prothrombin (II)



Thrombin (IIa)

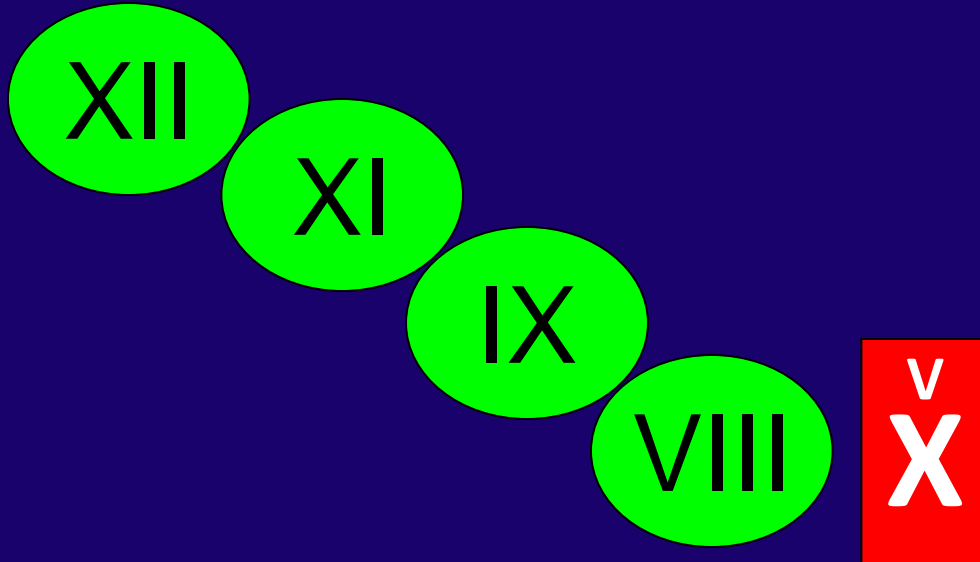
Fibrinogen



Fibrin

Coagulation made easy - the aPTT

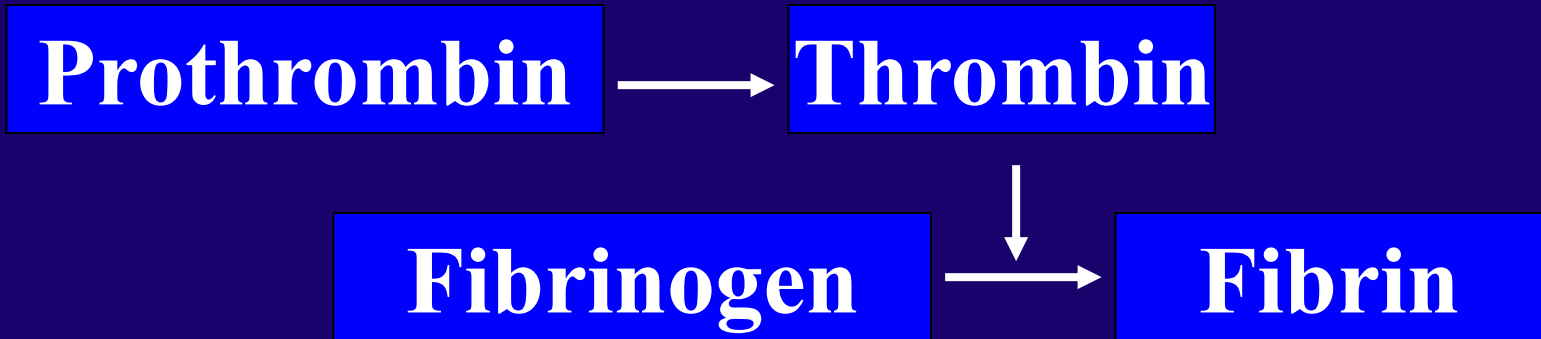
The PTT Pathway



The PTT pathway has all those hideous roman numerals. . .

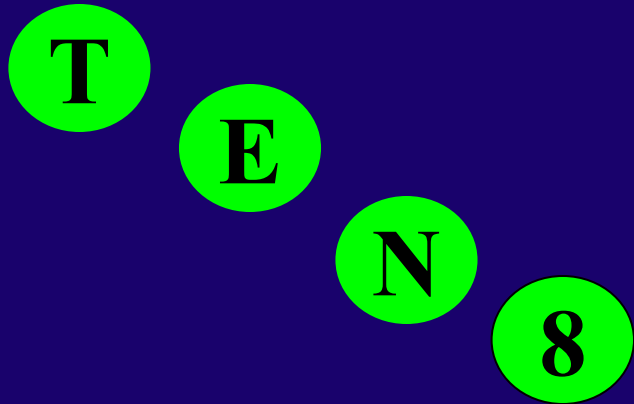
How are we going to remember them?

Hmmmmm.



Coagulation made easy - the aPTT

The PTT Pathway



Well, just remember that the PTT is a basic TENET (TEN8) of hematology. TENET stands for.

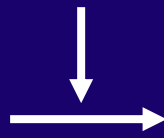


Prothrombin



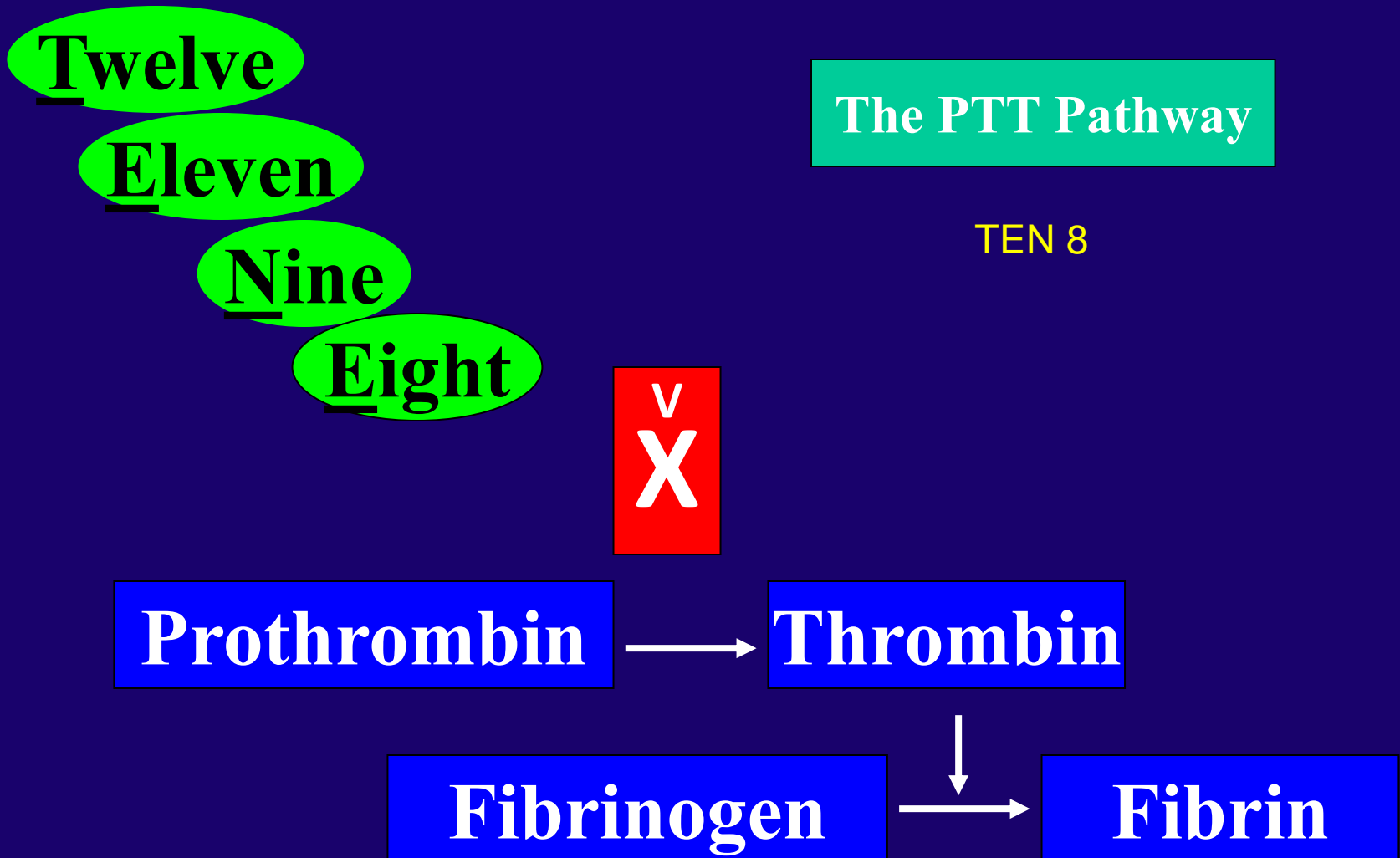
Thrombin

Fibrinogen

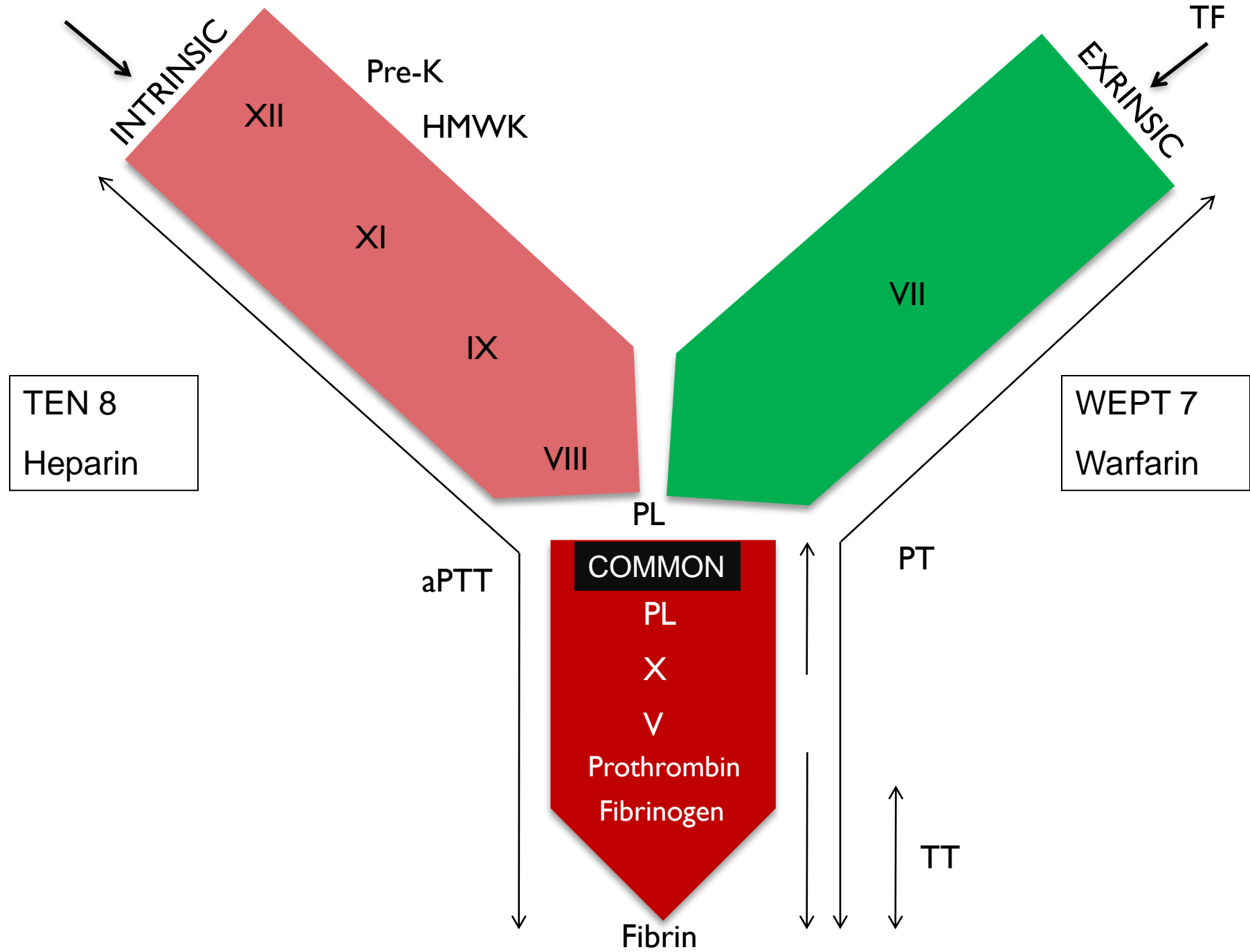


Fibrin

Coagulation made easy - the aPTT



CONTACT ACTIVATION



PT

Platelet Poor Plasma Sample (Containing all Factors)

+

PT Reagent

(Tissue Factor, PL, Ca⁺⁺)

Abnormality	Interpretation
Isolated Prolonged PT	Factor VII deficiency
Prolonged PT in association with other coagulation abnormalities	<p>Vitamin K deficiency</p> <p>Vitamin K antagonists e.g. warfarin, phenindione, rodenticides</p> <p>Liver disease</p> <p>Malabsorption (leading to vitamin K deficiency)</p> <p>High concentrations of unfractionated heparin</p> <p>Direct thrombin inhibitors e.g. Lepirudin, argatroban</p> <p>Afibrinogenaemia and dysfibrinogenemia</p> <p>Dilutional coagulopathy e.g. massive blood transfusion</p> <p>Multiple clotting factor deficiencies e.g. FV and FVIII deficiency</p> <p>Abnormalities of the vitamin K cycle e.g. mutations within the <i>VKORC1</i> gene</p>

aPTT

Platelet Poor Plasma Sample (Containing all Factors)

+

aPTT Reagent

(PL, Silica)



Incubation

+

Calcium Chloride

Abnormality	Interpretation
Isolated Prolonged APTT	<p>Deficiencies of either XII, XI, IX & VIII.</p> <p>Acquired clotting factor inhibitors - these are most commonly directed against FVIII.</p> <p>Lupus anticoagulant [LA]</p>
Prolonged APTT + Prolonged PT	<p>Vitamin K deficiency</p> <p>Liver disease</p> <p>Direct thrombin inhibitors including Hirudin, Argatroban and Dabigatran.</p> <p>DIC - due to the consumption of clotting factors</p> <p>Massive blood transfusion leading to a dilutional coagulopathy</p>

All are components of hemostasis except-

- a) Blood Platelets
- b) Red blood cells
- b) Endothelial Cells
- c) Plasma Coagulation Factors

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Steps involved in hemostasis include all except-

- a) Vasodilatation
- b. Formation of a platelet plug
- c. Blood Coagulation
- d. Clot retraction and thrombus dissolution

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Which organ is primarily responsible for the formation of coagulation factors?

- a) Kidney
- b) lungs
- c) liver
- d) Brain

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Which test evaluates the extrinsic pathway?

- a). PT
- b). PTT
- c). TT
- d). Closure time
- e). Bleeding time

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- a). PT WEPT 7
- b). PTT
- c). TT
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Heparin is monitored by the following test-

- a). PT
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Which of the following these patients may have a normal PTT?

- a) Thrombocytopenia
- b) Hemophilia A
- c) Hemophilia B
- d) Patient on heparin

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**Prolongation of Bleeding time seen in the following disorders
except-**

- a) Thrombasthenia
- b) Thrombocytopenia
- c) Von Willebrand ds.
- d) Hemophilia

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Platelet poor plasma means platelet count less than---

- a) 50,000/cmm
- b) 10,000/cmm
- c) 1,00,000/cmm
- d) 15,000/cmm

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2/Female child – Easy Bruising

	Test Patient	Reference Range
PT	13s	11-14s
APTT	105s	23-35s
Fibrinogen(Clauss)	2.7g/L	1.5-4.0g/L
Thrombin Time	13s	10-13s

What Next

- Repeat Assay
- Mixing studies: The prolonged APTT corrects in a mix with normal plasma
- What factor assays would you request and why?

FVIII, IX , XI assays.

The FIX assay was normal but the FVIII assay was <5 IU/dl.

Thank You

