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Hemodynamic Disorders, Thromboembolic Disease and Shock (Part 1)

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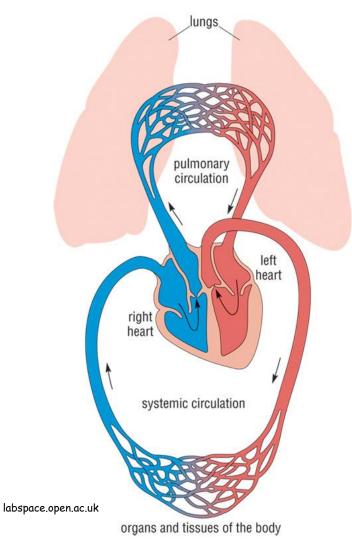
2nd Department of Pathology

Normal fluid homeostasis

- Vessel wall integrity
- Intravascular pressure and osmolarity in physiologic ranges
- Maintaining blood as a liquid

Protagonists

- Heart
- Blood vessels and lymphatic vessels
- Blood

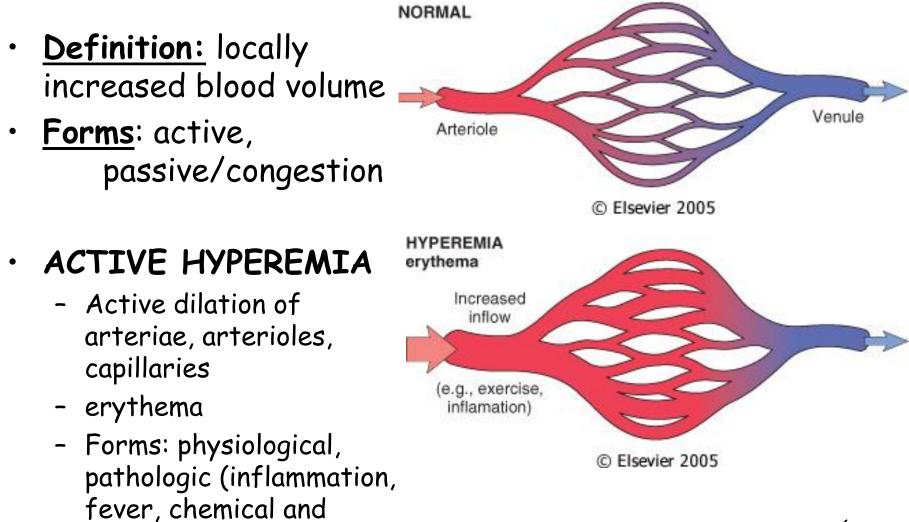


Hemodynamic disorders

- Hyperemia (active and passive)
- Edema
- Hemorrhage
- Thrombosis
- Embolism
- Infarction
- Shock

HYPERAEMIA

HYPEREMIA I.



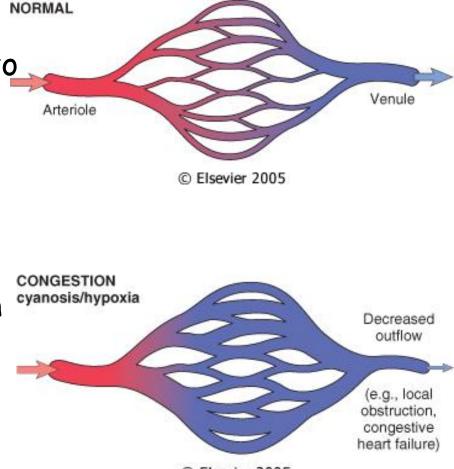
physical injury)

HYPEREMIA II.

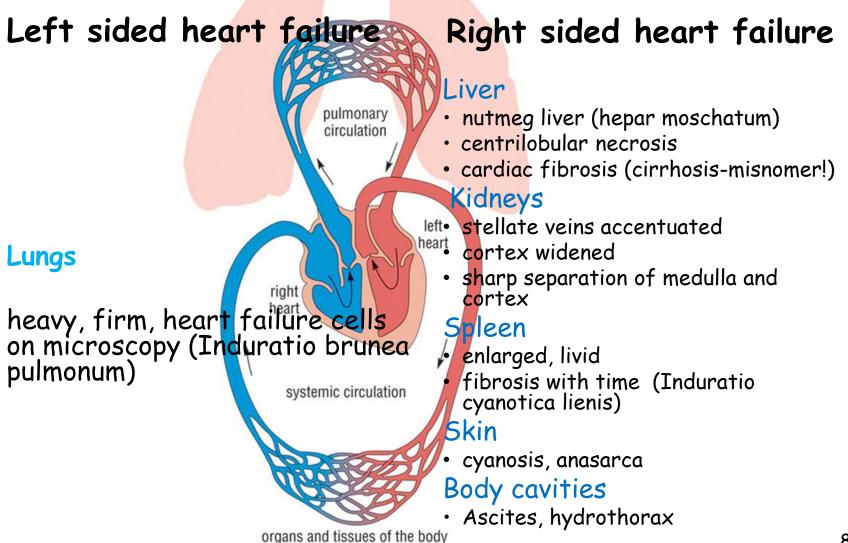
· PASSIVE HYPEREMIA

(congestion)

- dilation of venous side due to decreased outflow
- cyanosis, hypoxia
- <u>Causes</u>:
 - Systemic: congestive heart failure
 - Local: thrombosis, obstruction
 - Deep venous thrombosis-legs
 - Pylethrombosis- portal congestion
 - V.cava sup. syndrome



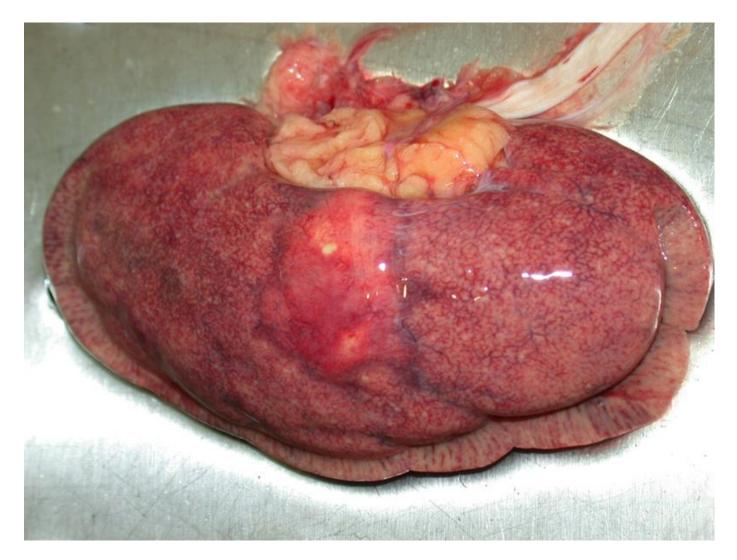
Consequences of systemic chronic congestion



Systemic congestion-Nutmeg liver



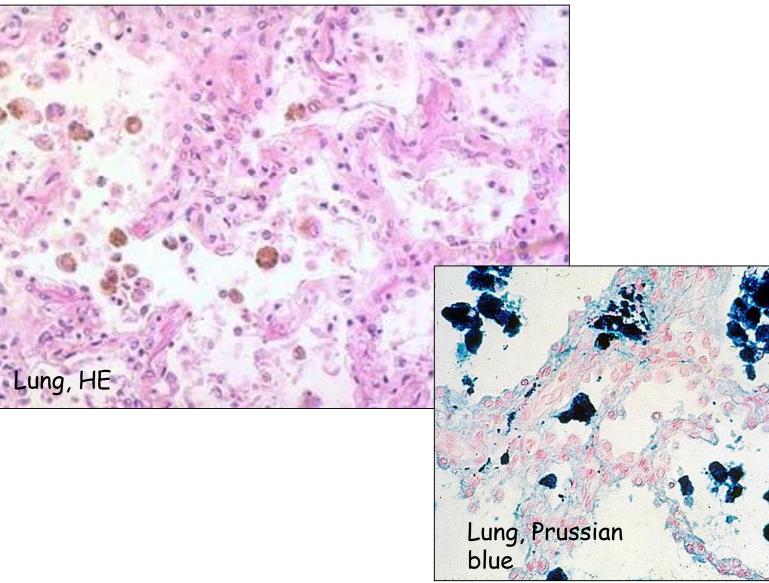
Systemic congestion (+?) - kidney



Lung, chronic passive hyperemia/ congestion



Hemosiderin in heart failure cells



Local chronic congestion

- May occur in every organ
 - E.g. Vena cava superior syndrome, Budd-Chiari sy (hepatic vein thrombosis), extremities etc.

Local congestion



EDEMA

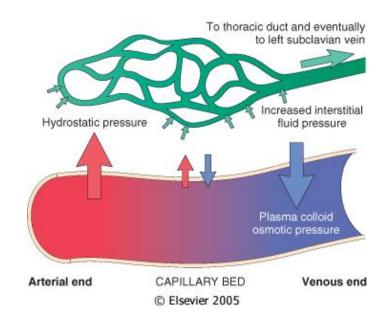
EDEMA

- <u>Definition</u>: increased fluid content in the interstitium (transudate or exudate)
- <u>Causes</u>:
 - Increased intravascular hydrostatic pressure
 - (arteriolar dilation or impaired venous return)
 - (local- eg. Deep venous thrombosis of the legs,
 - systemic- eg. Congestive heart failure)
 - Decreased plasma colloid osmotic pressure
 - (eg. nephrosis sy, cirrhosis, protein malnutrition)
 - Lymphatic obstruction (lymphedema, elephantiasis)
 - Salt and water retention (GN, Acute renal failure)
 - Inflammation (exudate)

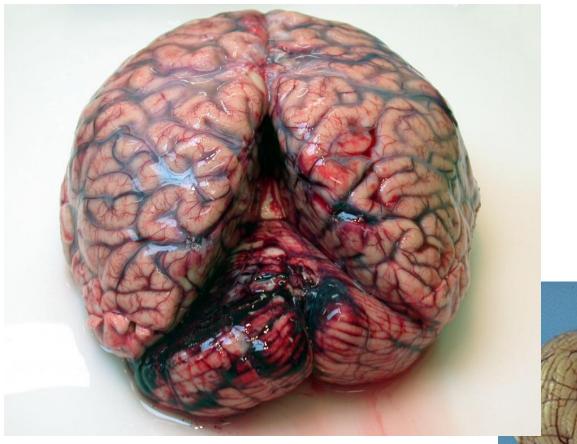
Normal fluid homeostasis:

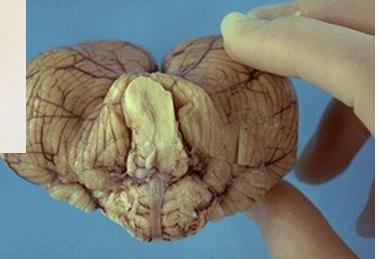
60% of body weight is water

2/3 intracellular, 5% blood, remainder within interstitium

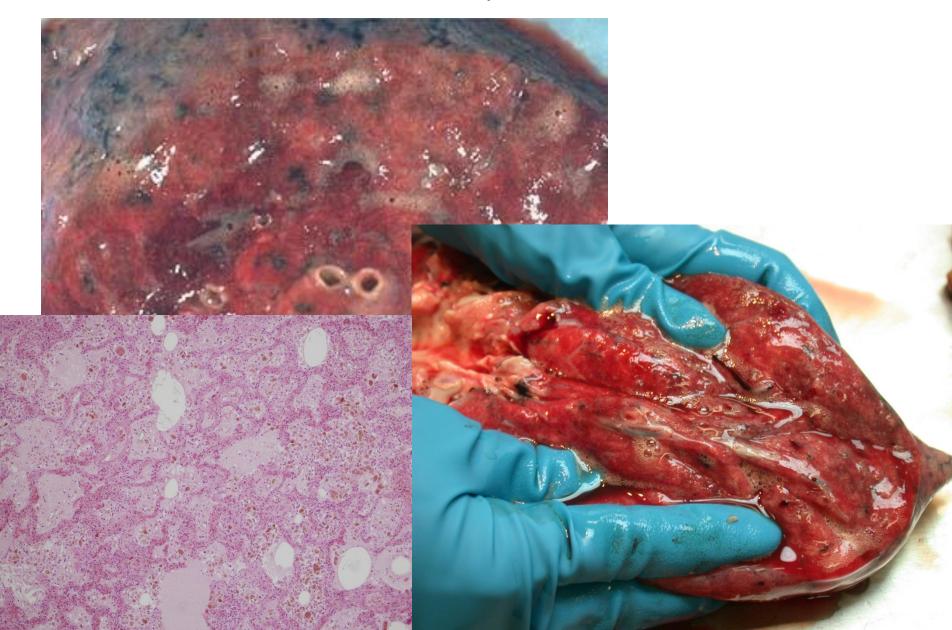


Cerebral edema- tonsillar herniation





Pulmonary edema











HEMORRHAGE

Hemostasis-protagonists

- Vascular wall (endothelium)
- Platelets
- Coagulation cascade

Normal hemostasis-sequence of events

- Arteriolar vasoconstriction (major regulator: endothelin)
- Platelet adherence, activation and aggregation > Primary hemostasis (major regulator: ECM)
- Fibrin meshwork creation, additional platelet recruitment Secondary hemostasis (major regulator: Tissue factor)

Hemorrhage

- <u>Definition</u>: Extravasation of blood
- Pathogenesis:
 - Rupture of vessel wall (haemorrhagia per rhexim)
 - Erosion of vessel wall (haemorrhagia per arrosionem)
 - Vascular wall disturbances and other hemorrhagic diatheses (haemorrhagia per diapedesim)
 - Vessel wall abnormalities: due to hypoxia, infections, drugs, impaired collagen synthesis, Henoch-Schönlein purpura, Hereditary hemorrhagic teleangiectasia etc.

• Other hemorrhagic diatheses

-Thrombocytopenia (low platelet count)

Decreased platelet production

Bone marrow diseases, bone marrow infiltration, drug induced (Heparin-induced thrombocytopenia), infections (HIV associated!) etc.

Decreased platelet survival

- Immune thrombocytopenic purpura (ITP, autoimmune)
- Thrombotic microangiopathies (TTP: thrombotic thrombocytopenic purpura, HUS: Hemolytic- uremic syndrome)

-Thrombasthenia (defective platelet function): primary, secondary (aspirin!!!)

-Abnormalities in clotting factors

- Primary, or Secondary (acquired eg. In hepatic diseases!)
- Von Willebrand disease
- e Hemophilia A (factor VIII deficiency)
- e Hemophilia B (Factor IX deficiency- Christmas disease)
- e Hemophilia C (Factor XI deficiency)

Disseminated intravascular coagulation (DIC, consumption coagulopathy)

» Causes: obstetric complications, infections, neoplasms, excessive tissue injury

23

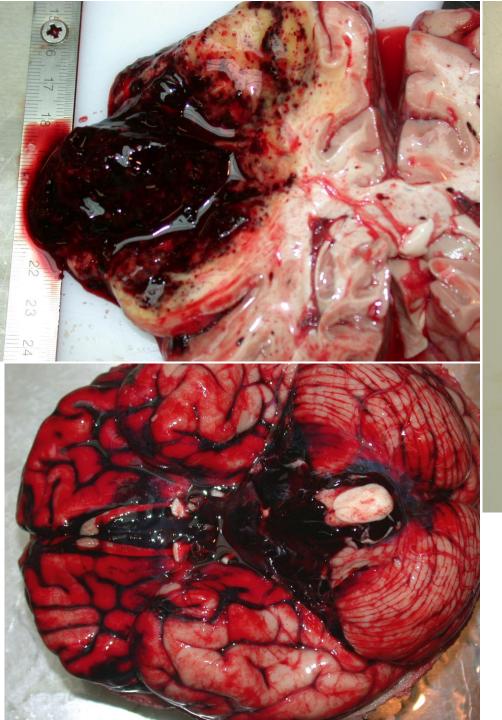
» Hemorrhage and thrombosis

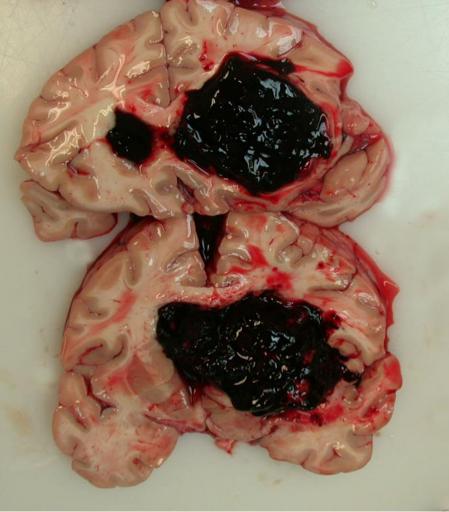
Categories

- By extent
 - Exsanguination, hematoma, suffusion, ecchymosis, petechia, purpura
- By localization

eg: hemothorax, hemopericardium, hemascos, hemarthrosis, epistaxis, hematemesis, melena, hematochesia...

By pathomechanism...





Haemorrhagia per rhexim (by rupture)- cerebral

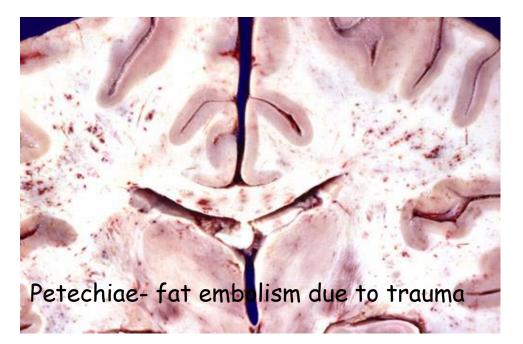


Haemorrhagia per arrosionem (by erosion)

(hypopharynx tumor)

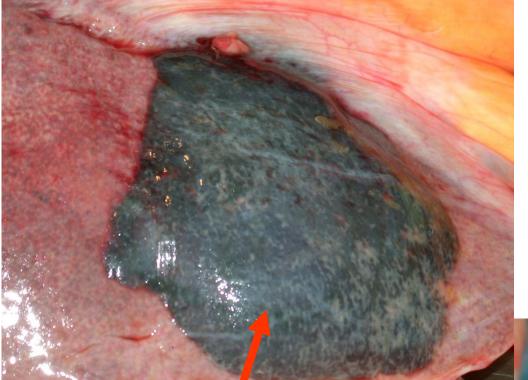






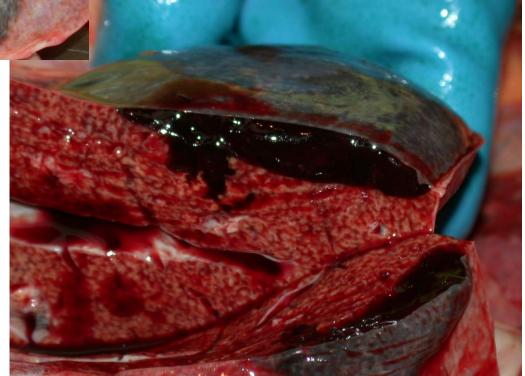
Haemorrhagia per diapedesim (by diffusion)

Categories by extent...



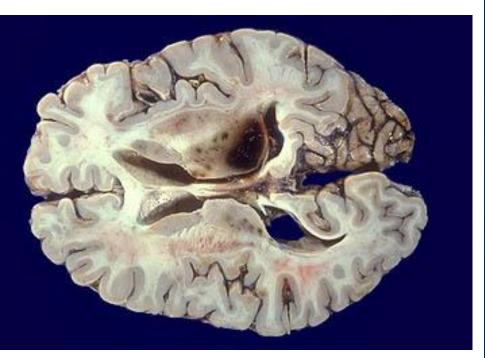
Subcapsular hematoma – liver

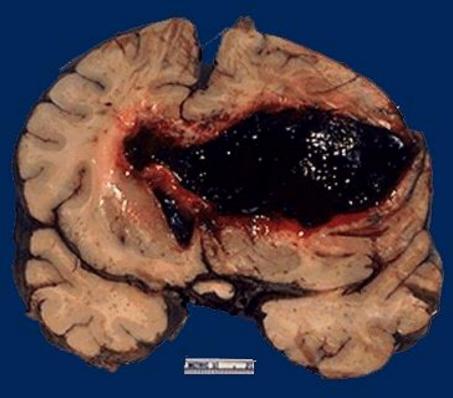
hematoma



Suffusion, hematoma

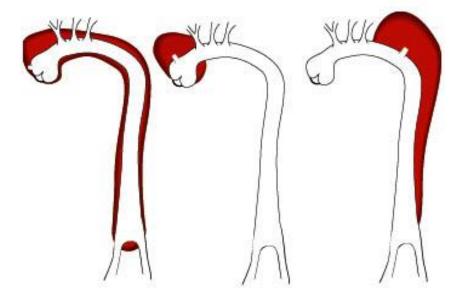






Thalamus hemorrhagehypertension

Apoplexia cerebrihypertension





Hemopericardium- due 33 to aortic dissection

THROMBOSIS

THROMBOSIS

- <u>Definition</u>: coagulation of blood within vessels or heart chambers
- <u>Pathogenesis</u> (Virchow's triad)
 - 1. Endothelial injury (inflammation, hypertension, atherosclerosis etc.)
 - 2. Stasis or turbulence of blood flow
 - 3. Hypercoagulability
 - Primary (Genetic predisposition) (e.g. Leydenmutation in factor V. (resistant to cleavage by protein C)
 - Secondary (Acquired): smoking, obesity, drugs, pregnancy, malignant tumors (e.g. pancreas cc- Trousseau phenomenon), antiphospholipid antibody syndrome ³⁵

Classification by localization 1.

Arterial thrombosis

Pathogenesis: endothelial injury, turbulent blood flow (due to atherosclerosis, vasculitis)

Complications:

- Ischemia (non occlusive thrombus)
- infarction (occlusive thrombus)

a.coron.-AMI, angina pectoris Cerebral arteries- TIA, stroke, status lacunaris a. mesenterica sup. or inf.- bowel infarction

Classification by localization 2. <u>Venous thrombosis</u> (phlebothrombosis)

Pathogenesis: stasis (varicositas, immobilization) thrombophlebitis

Localization: 90% legs

Periprostatic, parametrial plexus, dura sinuses, v. portae, vv.hepaticae, Trousseau phenomenon: migratory thrombophlebitis

Complications: Ulcus cruris

Postthrombotic sy Vena cava superior sy Vena cava inferior sy- congestion in lower extremities, pelvis

Classification by localization 3.

Thrombosis in heart chambers (like arterial thrombosis)

Causes: endothelial injury (eg. AMI), turbulent blood flow (dilatation, AMI)

Complications: atrial, ventricular, valvular (endocarditis maranthica) thrombi may be the source of **embolism**

Fibrin (Hyalin) thrombi

- In arterioles, capillaries, venules (disordered microcirculation)
- Composed of platelets, fibrin
- · DIC

Secondary hypercoagulable states High risk for thrombosis

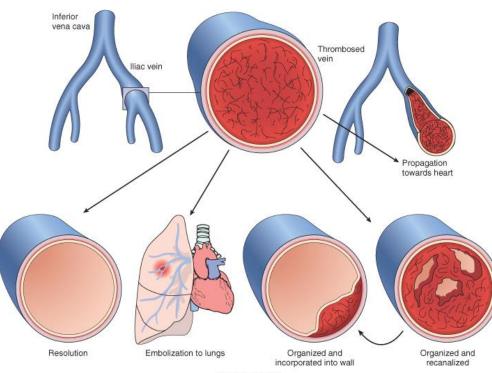
- Prolonged bed rest or immobilization
- \cdot Myocardial infarction
- Atrial fibrillation
- Tissue damage
 - (including surgery fractures, burns etc)

- Prosthetic cardiac valves
- Disseminated intravascular coagulation
- Heparin-induced thrombocytopenia
 HIT (unfractionated heparin)
- Cancer
- Antiphospholipid antibody syndrome
 - •("lupus anticoag.sy")

Secondary hypercoagulable states Lower risk for thrombosis

- Cardiomyopathy
- Nephrotic syndrome
- Hyperestrogenic states (Pregnancy,postpartum)
- Oral contraceptives
- Hyperlipidaemia
- Sickle cell anaemia
- Smoking

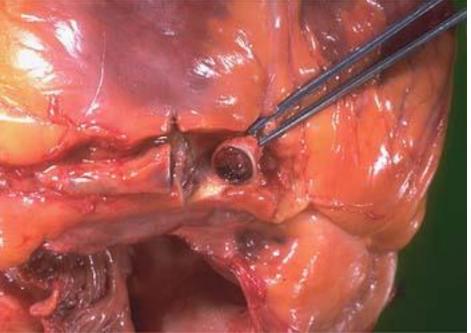
FATE OF THROMBI



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- Resolution
- Propagation
- Embolization
- Organization, recanalization

Arterial thrombus



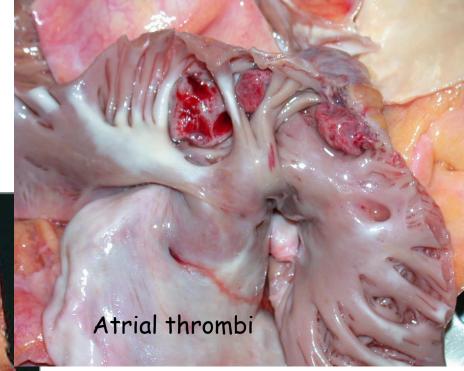
a. coronaria thrombosis

Venous thrombosis



Thrombus in iliac vein

Cardiac thrombi



Endocarditis (thrombotic vegetations)

Mural thrombus-left ventricle

EMBOLISM

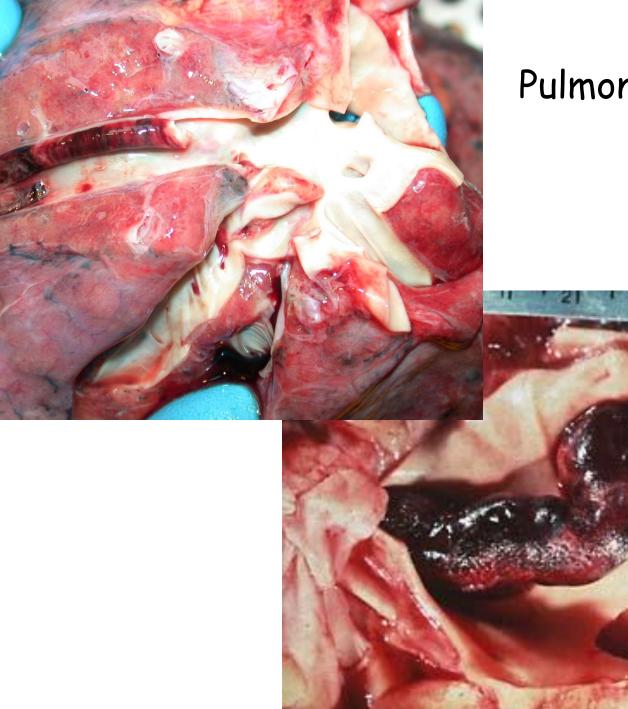
- <u>Embolus</u>: detached intravascular gaseous, liquid or solid mass carried by the bloodstream from its site of origin to another site where it causes vascular obstruction and subsequent tissue damage (necrosis)
- Forms:
 - Thromboembolism
 - Fat embolism
 - Air embolism
 - Amniotic fluid embolism-pulmonary edema, ARDS, DIC
 - Cholesterol embolism- kidney
 - Bacterial embolism- e.g. infective endocarditis
 - Foreign body embolism- i.v. drug abusers

Thromboembolism 1.

<u>Pulmonary thromboembolism</u>

Source: deep venous thrombosis of lower extremities Possible complications: - sudden death (saddle embolus)

- pulmonary hypertension, cor pulmonale
- hemorrhage, hemorrhagic infarction



Pulmonary embolism

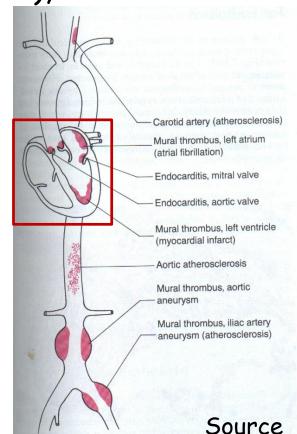
Thromboembolism 2.

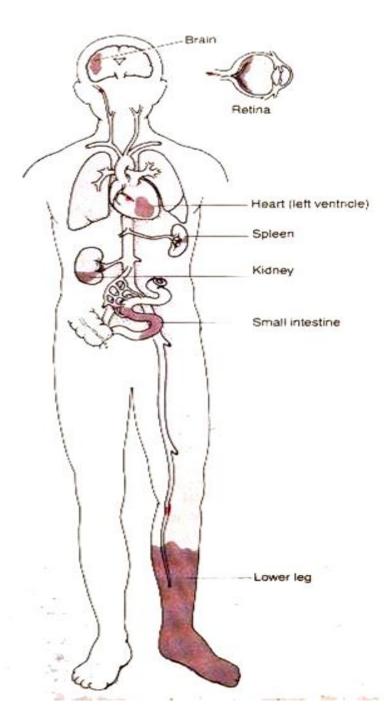
Systemic thromboembolism

Source: -80% mural thrombi within heart chambers (AMI, left atrial thrombi) -others: aortic atherosclerosis, aortic aneurysm, vegetations (in endocarditis), unknown origin Paradoxical embolism!

Complications: arterial embolization -in lower extremities (75%) -in the brain (10%), -in arteries of bowel, kidney, spleen, upper extremities

 \rightarrow INFARCTION





Systemic arterial embolization

lower extremities brain retina heart spleen kidney small bowel

Fat embolism

- After fractures of long bones
- Clinical signs: respiratory distress, neurologic symptoms (restlessness, irritability, delirium, coma), anemia, thrombocytopenia
- Mechanic and toxic injury

Air embolism

Causes:-Trauma (chest wall injury) -obstetric complications -decompression disease (eg. Scuba divers) -caisson's disease Complications: neurological symptoms,

pulmonary hemorrhage and edema, atelectasis, epiphyseal necrosis of long bones