Hemodynamic Disorder and cardiovascular diseases

台北醫學大學 病理學科 林永和 kevinyhl@tmu.edu.tw

(A) Hemodynamic Disorder

- Edema
- hyperemia or congestion
- hemorrhage
- thrombosis
- embolism
- infarction
- shock

Edema

- an excess of fluid in the interstitial spaces and/or body cavity
- 1. localized pitting edema :acute inflammation, allergy, venous obstruction, lymphatic edema
- 2 tissue or organ periorbital area, pulmonary edema, brain edema; ascites, hydrothorax (pleural effusion), pericardial effusion
- 3 generalized -- anasarca: heart, kidney, starvation
- transudate, exudate(more cell and protein)

Inducing reasons of edema

A) increased hydrostatic pressure (blood vessel) 1).venous return pressure increase 2) arterial dilation EX: deep vein thrombosis, congestive heart failure (backward heart failure) B) decreased oncotic pressure (plasma) 1). Hypoproteinemia. 2). Hypoalbuminemia EX: nephrotic syndrome, cirrhosis C) obstruction of the interstitial fluid drainage (lymphedema) EX: tumor, surgery, radiation, filariasis D) increased tissue hydration (sodium retention) EX: hyperaldosteronism

E) inflammation: congestion, vascular permeability increase

clinical signs of edema

- variable
- dependent on causes, rate, severity EX: lung and brain edema

Hyperemia

- 1. active (artery, capillary) \rightarrow arterial and capillary dilatation \rightarrow inflammation
- 2. passive (congestion), vein return obstructiona) acute
 - local abrupt vein return obstruction, thrombosis, organ torsion

systemic – acute heart failure, asphyxia

 b) chronic → ischemia → cell degeneration or death local – long term, progressive vein obstruction systemic – chronic heart failure

Hemorrhage – extravasation of blood

- *acute or chronic;
- *short term or long term;
- *external or internal
 - 1. types of hemorrhage
 - a) petechia: 1-2 mm mucosa, skin
 - vascular pressure increase, platelet decrease
 - b) purpura: $\geq 3mm skin$
 - trauma, vasculitis, increase in vascular fragility
 - c) ecchymosis: ≥ 1 cm skin
 - trauma

hemorrhage

Causes : trauma, vasculitis, tumor, hypertension, aneurysm rupture, hemorrhagic diathesis **Clinical terms:**

• hemoptysis (喀血)

- hematemesis (吐血) hematochezia (便血)
- hematuria (血尿)

- melena (瀝青性便血)
- metrorrhagia (陰道出血)
- menorrhagia (經血過多) hematoma (血腫血塊)

Thrombosis

- clotting, is transformation of the fluid blood into a solid aggregate encompassing blood cells and fibrin.
- Pathogenesis: endothelial cell injury → platelet aggregation and deposition → hemostatic plug formation → vascular statis
- 2) Factors of intravascular coagulation:
 - a) coagulation proteins b) platelets c) endothelial cells
- 3) Virchow's triad: three predisposing conditions for formation of pathologic thrombi
 - a) endothelial cell injury
 - b) hemodynamic change
 - c) hypercoagulability of the blood

Thrombosis

4) Types

- a) arterial thrombosis \rightarrow heart, large artery \rightarrow multiple organs infarct
- b) venous thrombosis \rightarrow vascular stasis leg vein \rightarrow local congestion edema, lung infarct
- 5) Fate of the thrombi:
 - a) Small thrombi lysis
 - b) Large thrombi :1) narrowing, occlusion
 - 2) organization –recanalization
 - 3) emboli ischemic infarction
- *clinical symptoms of thrombosis: 1) size 2) extent 3) rapid
 - a) heart, cerebral artery \rightarrow death
 - b) pulmonary Emboli→high mortality
 - (often not recognized clinically)

Disseminated intravascular coagulation

(DIC)

pathogenesis: activation and aggregation of intravascular fibrin protein \rightarrow consumption of platelet and coagulation factor \rightarrow resulting in hemorrhagic change

reason: infection, amniotic fluid, malignant tumor,

or unknown cause

clinical signs:

- diffuse microthrombosis formation → ischemia of brain, lung, heart, kidney
- 2) consumption of platelet and coagulation factor \rightarrow hemorrhage
- * injury of red blood cell when pass through the fibrin thrombosis → lysis → microangiopathic anemia

Embolus and Embolism

- a detached intravascular solid, liquid or gaseous mass
- a procedure of an embolus is carried by blood to a distal site fixed in the vascular lumen
- Forms of emboli:
- a) Thromboemboli—the most pattern
 - 1) pulmonary thromboembolism venous embolism
 - deep vein of leg and pelvis
 - small emboli—lung infarct, lung hemorrhage, silent (60-80%)
 - large emboli—acute lung failure, sudden death EX: traveler syndrome
 - 2) systemic thromboembolism arterial embolism
 - intracardiac mural thrombi (80%), dilated left atrium, atherosclerotic plaque, valvular vegetation, unknown origin (10-15%)
 - \rightarrow intestine, brain, spleen, kidney infarct

Forms of emboli

b) Liquid emboli -1) fat emboli, 2) amniotic fluid

1) long bone fracture (90%), fat tissue injury, burn

2) rare (1/50000) but deathful complication (20-40%) of labor, can't to prevent the amniotic content (amniotic fluid, fetal skin, hair, feces) into the mother vascular circulation \rightarrow lung infarct, DIC, hemorrhage and shock

c) Gaseous emboli (100cc volume)

Chest wall injury, decompression sickness, caisson disease—bone necrosis and organ infarct

d) Solid particle emboli - cholesterol crystal, tumor cells, bone marrow, foreign material

Infarction and infarct

A consequence of ischemic necrosis of distal tissue from an embolic event; an area of ischemic necrosis caused by occlusion of vessels

1) causes: thrombus or embolus (the most), vascular spasm,

compression, torsion, injury

2) Morphologic change:

Gross: triangle or wedge-shape, pale or hemorrhagic

Micro: coagulative necrosis, liquefactive necrosis, abscess (septic infarction)

3) type: a) white (anemic) infarct-arterial occlusion

 \rightarrow heart, spleen kidney, brain, solid organs

b) red or hemorrhagic infarct—venous obstruction or two sets

blood supply \rightarrow ovary, testis, intestine, lung, liver

The major factors that influence development of an infarct 1) nature of the vascular supply 2) rate of development of occlusion 3) vulnerability to hypoxia 4) oxygen content of blood

Shock (cardiovascular collapse)

Definition: a state of systemic hypoperfusion of tissue with blood 1) Type of shock:

a) cardiogenic shock --reduction in cardiac out—myocardial injury, ventricular occlusion, external compression b) hypovolemic shock --reduction in the effective circulating blood volume-hemorrhage, trauma, burn c) septic shock --loss of peripheral vascular tone—endotoxic shock, Gram (-), systemic infection *neurogenic shock—loss of sympathetic and vasomotor tone \rightarrow peripheral pooling of blood---pain, fear, spinal cord injury *anaphylactic shock—allergy \rightarrow IgE release, systemic vasodilation and \rightarrow increased vascular permeability—drug, nuts, insect bite

Symptoms and signs of shock

a) Skin, pale and cold

- b) Kidney, decreased GFR, oliguria, anuria, metabolic acidosis
- c) Acidosis potentiates heart failure
 - causes vasodilatation, peripheral pooling of blood
 - Left ventricular insufficiency raises the intrapulmonary venous pressure, pulmonary edema→shock lung, ARDS→respiratory acidosis
 - Anoxia cytokines release, (IL-1, TNF), vasodilatation

*anoxic endothelial cell \rightarrow lose their antithrombogenic properties \rightarrow revert into clot-forming cells, release clotting factors, combined with platelet, plasma clotting factors \Rightarrow DIC

Stages of shock

a) early or compensated shock: tachycardia, vasocontriction of peripheral arterioles, skin pale,

- * blood pressure is normal
- b) decompensated but reversible shock: tachypnea and shortness of breath (anoxia) acidosis, arterial dilatation, oligouria, confused

* hypotension

c) irreversible shock: circulatory collapse severe tissue and organ injury

Decreased blood pressure

Stimulate sympathetic nervous system a. thirst b.anxiety, restlessness Compensations c. tachycardia d.vasoconstriction, pallor to main of 2) Renin—angiotensin—aldosterone heart and brain a. vasoconstriction b. retention of Na+, function and water, oliguria Increased ADH secretion a. retention of water



pathologic changes of shock

- brain \rightarrow ischemic neuropathy
- heart→coagulation necrosis, subendocardial hemorrhage
- kidney \rightarrow acute tubular necrosis
- $lung \rightarrow diffuse alveolar damage (shock lung)$
- G-I system→hemorrhagic gastroenteropathy
- Liver \rightarrow fatty change
- Adrenal \rightarrow hemorrhagic necrosis

(B) Disease of cardiovascular system

1) Acquired heart disease:

Heart failure, Ischemical heart disease, Hypertensive heart disease, Cor pulmonale, Rheumatic heart disease, Infective endocarditis, Nonbacterial thrombotic endocarditis, Valvular heart disease, Myocardial disease,

2) Congenital heart disease,

3) Vascular diseases:

Arteriosclerosis, Varicose vein

Heart failure

A) right-sided heart failure:

- Cause: pulmonary artery occlusion, lung parenchyma disease, chronic inflammatory disease, lung tumor, chest wall deformity or exercise disturance
- Clinical signs: central venous pressure increase, hepatosplenomegaly, organ congestion, ascites, peripheral edema, nutmeg liver
- B) Left-sided Heart Failure:
 - Causes: hypofunction of left ventricle, aortic or mitral valve pathologic change
 - Clinical signs: lung edema, pleural effusion, cardiogenic asthma, hemoptysis, dyspnea

Ischemic heart disease

- causes: coronary arterial occlusion or spasm, vasculitis, tumor, hypovolemic shock, ventricular fibrillation → coronary artery blood flow decreased or interruption
- 2) clinical signs: angina pectoris, acute myocardial infarct, chronic ischemia heart disease, cardiogenic sudden death
- 3) gross and microscopic changes: myocardial necrosis, hemorrhage and fibrosis
- 4) complications: arrhythmia, left-sided heart failure, papillary muscle rupture, mural thrombus formation, heart rupture, ventricular aneurysm, pericarditis
- 5) death: arrhythmia, cardiogenic shock, thromboemboli

Hypertensive heart disease
1) definition : blood pressure >140/90mmHg
2) gross: left ventricle hypertrophy, dilatation, heart failure

3) side effects: pathological change in central nervous system and kidneys

Cor pulmonale

 cause: pulmonary hypertension, pulmonary thromboemboli, chronic obstructive lung disease, lung tumor, chest wall deformity or exercise disturbance

2) gross:

acute form – right ventricle dilatation, right-sided heart failure chronic form – right ventricle hypertrophy, right-sided heart failure

Rheumatic heart disease

 acute rheumatic fever – Group A beta, hemolytic streptococcus → pharyngitis

clinical signs:

- * myocarditis Aschoff's nodule (diagnostic characteristic),
- * migratory polyarthritis,
- * subcutaneous nodule,
- * erythema marginatum,
- * Sydenham's chores

#ASLO (antistreptolysin O) antibody elevation, fever Endocardium – vegetation, heart failure

Rheumatic heart disease

- 2) chronic rheumatic heart disease
- gross: valve deformity mitral & aortic valve
- \rightarrow atrophy, hypertrophy, fibrosis, calcification
- clinical signs: stenosis or insufficiency
 - \Rightarrow blood flow stasis, left ventricle dilatation,
 - mural thrombus formation, atrial fibrillation, lung congestion, heart failure

Infective endocarditis

- a) acute: present in healthy heart
 - *pathogen staphylococcus, pneumococcus sepsis bacterial vegetation – infective emboili, pyemia abscess
- b) subacute: present in heart with congenital deformity or valvular abnormality

*pathogen – streptococcus viridans bacterial vegetation, anemia, multiple emboli, splenomegaly

Nonbacterial thrombotic endocarditis

patient with chronic disease, or malignant tumor

gross: small nonbacterial vegetation over the leaflets of mitral valve or tricuspid valve

valvular heart disease

1) mitral valve stenosis

cause: congenital, rheumatic heart disease
clinical signs: progressive left heart failure, heart
murmurs, lung congestion and edema, pleural effusion
2) mitral valve insufficiency

cause: congenital, rheumatic heart disease, myocardial infarct

clinical signs: left ventricular hypertrophy, heart murmur

valvular heart disease

 3) aortic valve stenosis
 cause: congenital, rheumatic heart disease, myocardial hypertrophy, calcification
 clinical signs: left ventricle hypertrophy, heart
 murmur

4) aortic valve: insufficiency cause: congenital, syphilic aortitis
5) pulmonary valve stenosis cause: congenital

Myocardial disease: myocarditis

- Inflammation with myocardial necrosis
- (not induce by ischemia)
- a) cause: virus, bacterial, parasite, fungus, drug, alcohol, radiation
- b) clinical signs: abrupt or progressive heart failure
- c) gross or micro changes: cardiomegaly, myocardial necrosis
- d) complication: acute heart failure, arrhythmia, death

cardiomyopathy

unknown cause, (not by coronary artery disease, hypertension, valvular disease)

a) dilated cardiomyopathy

cause: unknown, may be associated with myocarditis or mutation of gene of myocardial contraction protein clinical signs: progressive heart failure

gross: heart enlarged and weight, dilatation of the four chambers, mitral and tricuspid insufficiency

micro: cardial cell hypertrophy, hyperchromatic nucleus, interstitial fibrosis

cardiomyopathy

b) hypertrophic cardiomyopathy

- cause unknown: but 25% associated with abnormal gene in cardial contractile protein
- clinical signs: progressive heart failure
- gross and micro changes: ventricle and septum hypertrophy, cardial cell hypertrophy and disarrayed distribution

cardiomyopathy

- 3) restrictive cardiomyopathy
 - ventricular space decrease
 - cause unknown
 - amyloidosis, glycogen storage disease, hemochromatosis, endocardial fibrosis, endocardial fibroelastosis

Congenital heart disease

- a) cyanotic
 - 1) tetralogy of Fallot (TOF)
 - 2) transition of great vessel (TFA)
 - 3) truncus arteriosus

Tetralogy of Fallot

- •Complex congenital defect of heart and major vessel
- •Cyanosis in newborn, 10% of CHD
- •Four typical lesions
- a) valvular stenosis (narrowing)
 - of the pulmonary artery
- b) ventricular septal defect
- c) dextroposition of the aorta
- d) hypertrophy of the right ventricle
- •Blue baby
- •Treat: surgical correction

Congenital heart disease

b) non-cyanotic

- 1) ventricular septal defect (VSD) the most
- 2) patent ductus arteriosus (PDA)
- 3) atrial septal defect (ASD)
- 4) coarctation of aorta
- 5) pulmonary stenosis (PA)
- 6) aortic stenosis (AS)

Septal defects: 30~40%

- a) Atrial foramen ovale
- b) Ventricle 1) foramen ovale (upper part)

2) muscular septa (lower part)

- VSD the most common CHD
 - L to R shunt
 - R ventricular hypertrophy
 - pulmonary hypertension
 - R to L shunt, cyanosis
- *easy detection by heart murmurs
- *defect small one, close spontaneous

large one, surgery

vascular disease

Arteriosclerosis

- 1) arteriolosclerosis
- muscular wall thickened, fibrinoid degeneration
- benign or malignant hypertension
- hypertensive heart disease, hemorrhagic stroke, hypertensive renopathy

2) atherosclerosis

- a) high risk factors: hyperlipiemia, hypertension, diabetes mellitus (DM), smoking
 - factors can not be changed: age, sex, hereditary
 - factors can be changed:
 - 1) lipid metabolism-related factors
 - diet, hyperlipidemia, obesity, D.M.
 - 2) hypertension
 - 3) clotting factors
 - 4) cigarette smoking

5) behavior

2) atherosclerosis

- b) muscular wall thick, lipid aggregation, fibrosis, calcification, hemorrhage, ulceration, thrombus formation, lumen narrowing, chronic ischemia
- c) aorta, coronary artery, cerebral artery, femoral artery, popliteal artery, radial artery
- d) complications: thromboemboli formation, aneurysm, myocardial infarct, ischemic stroke, peripheral infarct and necrosis

Arteritis

- 1) infective arteritis
- 2) syphilitic arteritis ascending aorta
- 3) thromboangiitis obliterans
- 4) giant cell arteritis

Aneurysm

abnormal vascular dilatation due to vascular wall destruction

- cause: atherosclerosis, trauma, inflammation, vascular degeneration, congenital
- gross change: compression, thrombus formation, hemorrhage
- type: atherosclerostic aneurysm, aortic dissection, mycotic aneurysm

diseases of vein

1) varicose vein

- cause: intraluminal pressure increase, extraluminal supporting decrease, vascular wall weakness
- common site: lower leg, esophagus, anus, umbilicus area
- complications: ulceration, hemorrhage, pigment deposition, abscess, fistula, thromboemboli formation
- 2) thrombophlebitis thrombotic occlusion of vein