#### **Regional Circulations** (pulmonary, skin, muscle, cerebral, splanchnic, renal, fetal)

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- Blood flow through lungs is virtually equal to the blood flow through all other organs.
- Functions:
  - provide the gas exchange
  - blood reservoir
  - mechanical, chemical and immunological filter

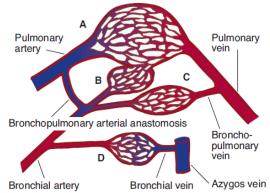


- Arteries (differences compared to the arteries in the systemic circulation)
  - bigger total cross-section of all pulmonary arteries
  - smaller thickness of the vessel walls
  - high compliance
- Capillaries
  - wide, abundant anastomoses form a net surrounding alveoles
  - time of passage, area of perfused capillaries at rest and intensive exertion
- Veins
  - high compliance (blood reservoir, ortopnoe)

Blood pressure in pulmonary circulation



- Nutrient circulation
  - physiological arteriovenous shunt



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#### Lymphatic vessels

- fast transport of proteins and various particles from the peribronchial and perivascular tissue  $\rightarrow \downarrow$  formation of the tissue fluid ~ prevention of the pulmonary edema

Minimal filtration in pulmonary capillaries physiologically!

- 1. pressures in intersticium and pulmonary capillaries
- 2. permeability of pulmonary capillaries

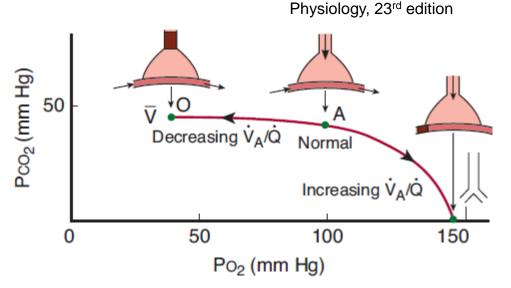


- Regulation of blood flow in lungs
  - A. Systemic mechanisms
    - 1) Neural regulation (sympathicus, parasympathicus)
    - 2) Humoral regulation (circulating substances)
  - B. Local mechanisms
    - chemical (metabolic) autoregulation
       opposite reaction compared to systemic circulation (vasoconstriction)
  - C. Passive factors
    - cardiac output
    - gravity (blood distribution in lungs)



#### • Ratio of ventilation and perfusion

- kept constant (local metabolic autoregulation) non-ventilated alveolus - vasoconstriction
   non-perfused alveolus - bronchoconstriction
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- decreased ratio most often cause of hypoxic hypoxia in clinical practise (right-left shunt) → ↓ arterial blood saturation with O<sub>2</sub>



 content of CO<sub>2</sub> usually not changed (compensatory hyperventilation in other alveoles)





• Skin blood flow considerably varies (0.02-5 l/min).

#### **Function:**

- Metabolic demands of skin small (decubitus)
- Maintenance of body temperature

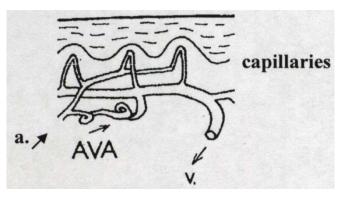
poikilothermic tissue

Arteriovenous anastomoses

- Protection against environment
- Maintenance of mean blood pressure



- Arteriovenous anastomoses
  - convoluted muscle vessels directly connecting arteriols and venules (low-resistance shunt)



Honzíková N - Poznámky k přednáškám z fysiologie (1992)

 regulated by sympathetic vasoconstrictive nerve fibers



- Regulation of skin blood flow:
  - Sympathetic nerve fibers
  - Humoral local factors (histamine, serotonine)

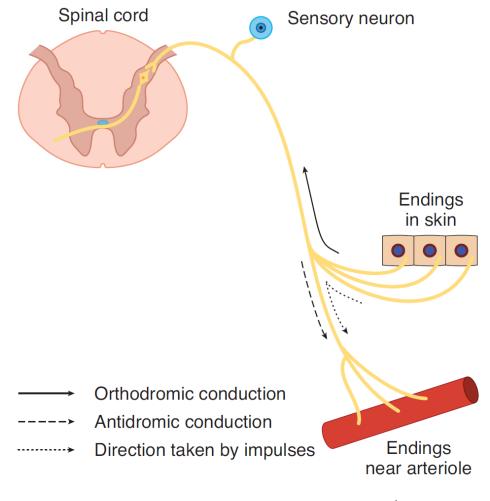


- Reaction on a temperature change:
  - 1) direct impact of a temperature change on the vessel tone
  - 2) excitation of skin thermoreceptors
  - 3) excitation of thermoreceptors in brain

reflex modulation of sympathetic vasoconstrictive activity



• Axon reflex



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#### **Muscle Circulation**



#### **Muscle Circulation**

- Function:
  - 1) Blood supply of muscles

the resting blood flow – 18% of the cardiac output vs. even 90% at intensive exertion (the local blood flow  $\uparrow$  even 20times)

2) Regulation of blood pressure

skeletal muscles – 40% of the body weight  $\rightarrow$  resistance of the muscle bloodstream has a high impact on the total peripheral resistance

 The blood flow during muscle activity is intermittent, during the tetanic contraction even zero (oxygen debt).

## **Muscle Circulation**

- Regulation of the muscle blood flow:
  - 1) Neural regulation

dominates at rest (vasocontriction through sympathicus - big dilation reserve)

2) Local chemical regulation

dominates at physical exertion (metabolic vasodilation)

almost linear increase of the flow with increasing metabolic activity

increased blood flow + increased O<sub>2</sub> extraction

 $\uparrow$  capillary pressure +  $\uparrow$  osmolarity  $\rightarrow$   $\uparrow$  filtration  $\rightarrow$  edema in active muscles





- provides:
  - 1) constant sufficient blood supply

(black-out during several seconds of the brain ischemia, irreversible damage during several minutes)

2) dynamic blood redistribution

(metabolic hyperaemia)



- Anatomical specialities of cerebral circulation:
  - *circulus arteriosus cerebri* (interconnection of main cerebral arteries by anastomoses)
  - 2) very high density of capillaries

(3000 – 4000 capillaries / mm<sup>2</sup> od the grey matter)

~ minimalization of diffuse distance for gases and other substances

3) very short arteriols

(almost 1/2 of the vasal resistance falls on arteries which are abundantly innervated)



- Functional adaptation of cerebral circulation:
  - 1) high and stable blood flow
  - 2) high O<sub>2</sub> extraction
  - 3) well developed autoregulation (myogenic and metabolic)
  - 4) high reactivity on changes of CO<sub>2</sub> concentration
  - 5) local vs. total hypoxia
  - 6) innervation



- Special physical conditions of cerebral circulation:
  - 1) solid cover of brain by skull

Monro-Kelli theory

 $\rightarrow$  flow may be increased only by acceleration of the blood flow, not by an increase of capacity of the bloodstream

 $\rightarrow$  Cushing reflex

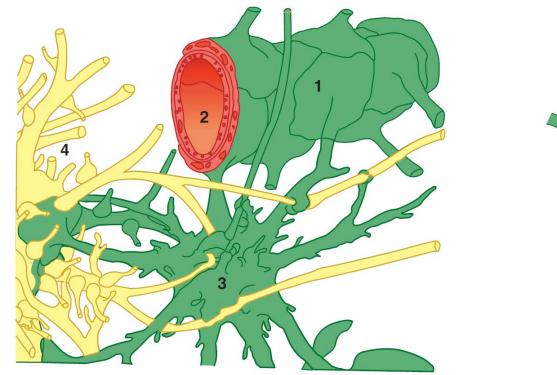
#### 2) gravity

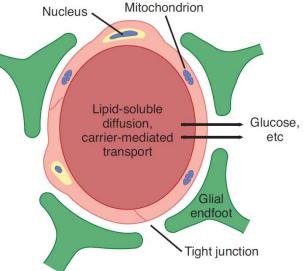
orthostatic reaction (postural syncope)



#### • Blood-brain barrier

cerebral capillaries - tight inter-endothelial connections





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#### Blood-brain barrier

#### By free diffusion:

 $\rightarrow$  lipophilic substances (O<sub>2</sub>, CO<sub>2</sub>, xenon; unbound forms of steroid hormones)

→ water (aquaporins; osmolality of blood and cerebrospinal fluid is identical!)

→ glucose – the main source of energy for neurons (free diffusion would be slow – accelerated by GLUT)

#### By transcellular transport (regulated):

 $\rightarrow$  ions (e.g. H<sup>+</sup>, HCO<sup>3-</sup> vs. CO<sub>2</sub> !)

 $\rightarrow$  transporters for thyroid hormones, some organic acids, choline, precursors of nucleic acids, aminoacids, ...

- Blood-brain barrier
- Functions:
  - maintenance of constant composition of the neuron environment
  - protection of brain against endogenic and exogenic toxins
  - prevention of loss of neurotransmitters to the bloodstream



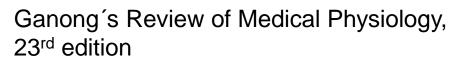
#### Cerebrospinal fluid

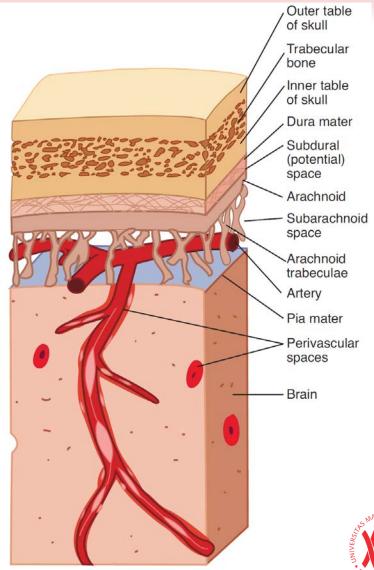
- localization
- composition
- volume ~150 ml, rate of production ~550 ml/d (exchange 3.7times/day)

Su	bstance	CSF	Plasma	Ratio CSF/Plasma
Na <sup>+</sup>	(meq/kg H <sub>2</sub> O)	147.0	150.0	0.98
K <sup>+</sup>	(meq/kg H <sub>2</sub> O)	2.9	4.6	0.62
$Mg^{2+}$	(meq/kg H <sub>2</sub> O)	2.2	1.6	1.39
Ca <sup>2+</sup>	(meq/kg H <sub>2</sub> O)	2.3	4.7	0.49
Cl⁻	(meq/kg H <sub>2</sub> O)	113.0	99.0	1.14
HCO3-	(meq/L)	25.1	24.8	1.01
Pco <sub>2</sub>	(mm Hg)	50.2	39.5	1.28
рН		7.33	7.40	
Osmolality	(mosm/kg H <sub>2</sub> O)	289.0	289.0	1.00
Protein	(mg/dL)	20.0	6000.0	0.003
Glucose	(mg/dL)	64.0	100.0	0.64
Inorganic P	(mg/dL)	3.4	4.7	0.73
Urea	(mg/dL)	12.0	15.0	0.80
Creatinine	(mg/dL)	1.5	1.2	1.25
Uric acid	(mg/dL)	1.5	5.0	0.30
Cholesterol	(mg/dL)	0.2	175.0	0.001

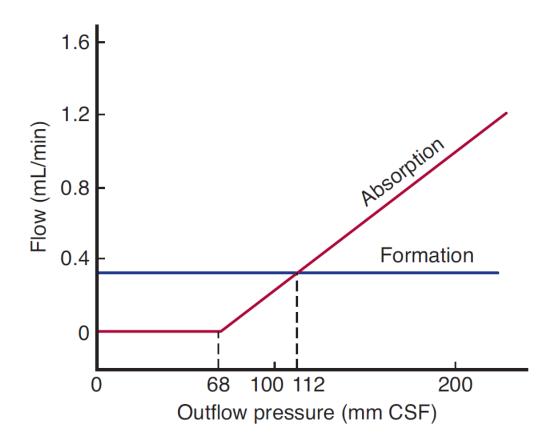
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- Cerebrospinal fluid Function:
  - protection of brain (together with menanges)





• Cerebrospinal fluid

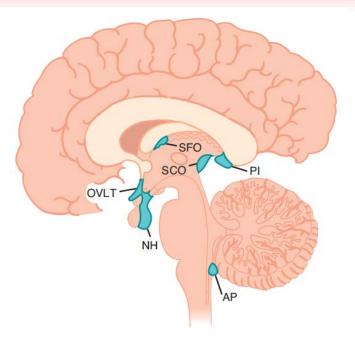


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# Paraventricular organs brain regions where the blood-brain barrier is missing (fenestrated capillaries)

- secretion of polypeptides (oxytocin, vasopressin, ...),
- chemoreceptive zones (AP)
- osmoreceptive zones (OVLT)



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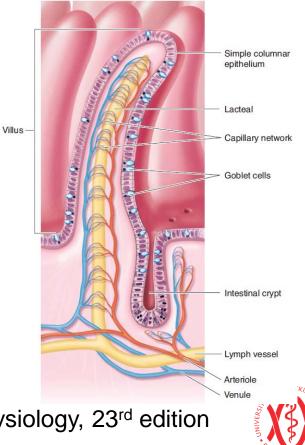
- blood flow through GIT including liver and pancreas
- blood flow through spleen
- Main functional roles:
  - metabolic function of GIT
  - blood reservoir
  - special (*e.g.* spleen removal and degradation of old/altered erythrocytes)



- Blood reservoir
- at rest ~20% of the total blood volume
- rich innervation with sympathetic vasoconstrictive fibers - α rec. (even 350 ml of the blood emptied into the systemic circulation during several minutes!)



- Intestinal circulation (a. coeliaca, a. mesenterica superior and inferior)
- submucous plexus, branches enter musculature and intestinal villi
- countercurrent exchange of substances



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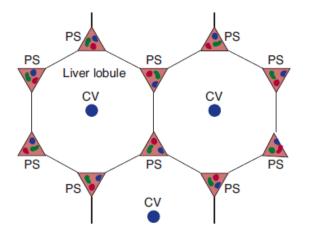
- Intestinal circulation (a. coeliaca, a. mesenterica superior and inferior)
- Regulation of blood flow:
  - metabolic vasodilation (mediators: adenosine,  $\downarrow$  [K<sup>+</sup>]<sub>e</sub> and  $\uparrow$  osmolarity)
  - neural regulation almost exclusively sympathicus,  $\alpha > \beta$  rec.  $\rightarrow$  vasoconstriction

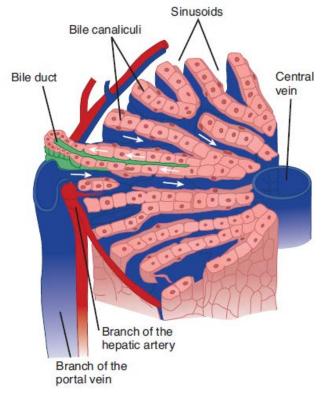


- Hepatic circulation (v. portae, a. hepatica)
- 25% of the cardiac output (~1.5 l/min)
  - $\frac{3}{4}$  v. portae,  $\frac{1}{4}$  a. hepatica Regarding O<sub>2</sub> supply, the ratio is opposite!
- portal circulation
  - 2 capillary bloodstreams in series (intestinal villi, liver sinusoids)
  - $\downarrow O_2$  content  $\rightarrow a$ . hepatica represents the nutritive hepatic circulation



- Hepatic circulation (v. portae, a. hepatica)
- functional unit acinus



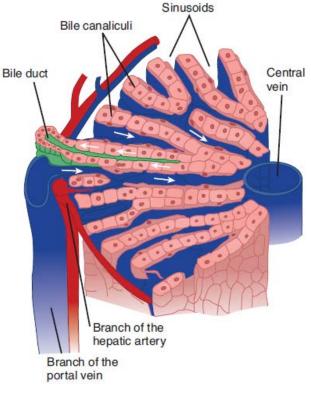


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- Hepatic circulation (v. portae, a. hepatica)
- pressures:
  - a. hepatica:
  - v. hepatica:
  - v. portae:
  - sinusoids:

90 mmHg 5 mmHg 10 mmHg 2.25 mmHg



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## **Splanchnic Circulation**

- Hepatic circulation (v. portae, a. hepatica)
- inverse regulation of blood flow in *v. portae* and *a. hepatica*:
  - <u>between meals</u>: many sinusoids collapsed, flow in *v*. *portae* low, adenosine formed constantly and washed less  $\rightarrow$  dilation of terminal hepatic arterioles
  - <u>after a meal:</u> flow in *v. portae* ↑, adenosine washed faster → constriction of hepatic arterioles, higher flow in *v. portae* opens so far collapsed sinusoids
- increased hepatic pressure (cirhosis)  $\rightarrow$  ascites



## **Splanchnic Circulation**

- Hepatic circulation (v. portae, a. hepatica)
- <u>Regulation of blood flow</u>:
  - neural: symp. vasoconstrictive fibers  $\alpha$  rec.
  - metabolic: adenosine  $\rightarrow$  vasodilation
  - passive: ↑ BP → passive dilation of portal vein radicles → ↑ liver blood amount

congestive heart failure

diffuse noradrenergic discharge due to  $\downarrow$  BP

• sufficient  $O_2$  supply is essential for liver function! -  $\downarrow$  flow  $\rightarrow \uparrow O_2$  extraction

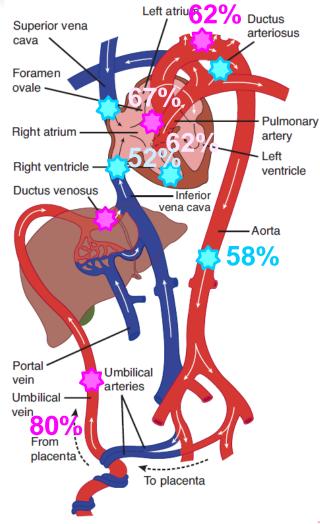
### **Splanchnic Circulation**

- Hepatic circulation (v. portae, a. hepatica)
- hepatic lymphatic circulation
  - formation of almost <sup>3</sup>/<sub>4</sub> of the body lymph
  - lymph rich on proteins (many plasmatic proteins are formed in hepatocytes + proteins from plasma due to the high permeability of sinusoids)





- placenta, umbilical vein
- liver, *ductus venosus*
- crista dividens, foramen ovale
- blood supply of the head and upper limbs
- v. cava superior and inferior
- the right ventricle
- ductus arteriosus
- aorta the blood supply of the lower part of body + 60% of the cardiac output is directed to placenta

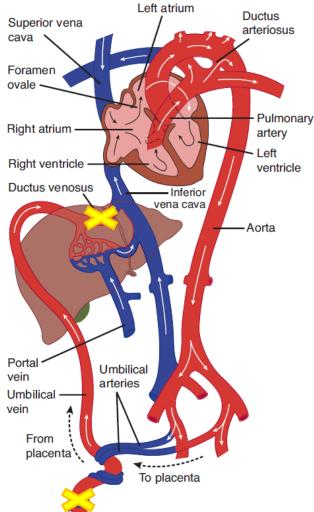




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- Changes after birth
- Closure of umbilical vein

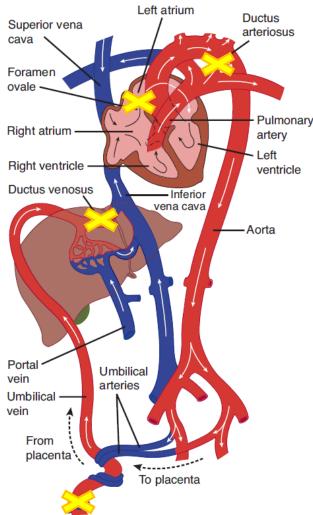
  - contraction of musculature of ductus venosus and its closure
- The first inspiration (due to asphyxia and cooling of the body)
  - ↓ resistance of the lung bloodstream
  - much more blood into lungs



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- Changes after birth
- Decrease of pressure in right atrium and its increase in left atrium due to:
  - filling of left atrium by the blood from lungs
  - ↓ venous return to right atrium due to closure of umbilical vein
  - left ventricle works against ↑ pressure in aorta
- Closure of formanen ovale
- Closure of ductus arteriosus



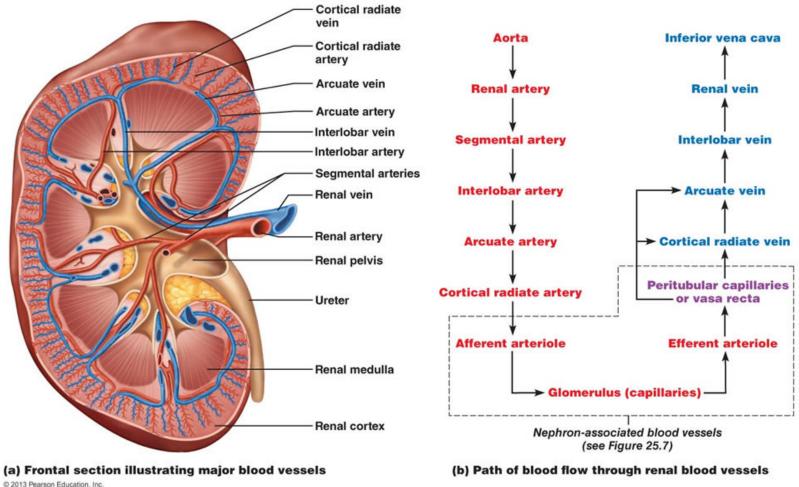
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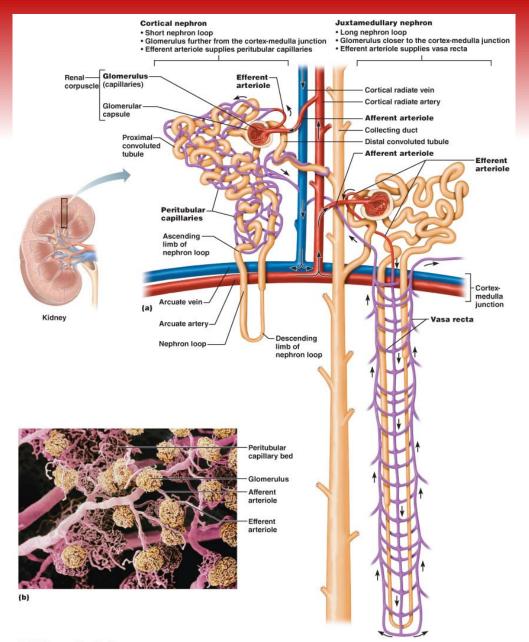
- main functions of kidneys
- High filtration rate requires an adequate blood supply!
  - kidneys form only ~0.4 % of the body weight
  - blood flow 1.2 l/min, ~25% of cardiac output
- distribution of blood flow is irregular, the most flows through cortex (glomeruli – filtration)







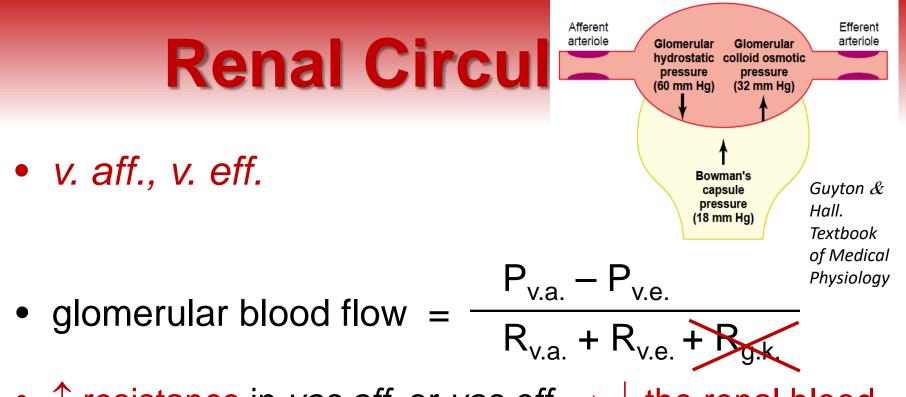
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- ↑ resistance in vas aff. or vas eff. → ↓ the renal blood flow (if the arterial pressure is stable)
- regulate the glomerular filtration pressure:

constriction of vas aff.  $\rightarrow \downarrow$  glomerular pressure  $\rightarrow \downarrow$  filtration constriction of vas eff.  $\rightarrow \uparrow$  glomerular pressure  $\rightarrow \uparrow$  filtration

- Regulation of renal blood flow:
  - 1) Myogenic autoregulation
  - 2) Neural regulation
  - 3) Humoral regulation



- Regulation of renal blood flow:
  - 1) Myogenic autoregulation
    - dominates
    - provides stable renal filtration activity by maintaining stable blood flow at varying systemic blood pressure



- Regulation of renal blood flow:
  - 2) Neural regulation
    - conformed to demands of systemic circulation
    - sympathetic system norepinephrine

light exertion/upright body posture  $\rightarrow \uparrow$  sympathetic tone  $\rightarrow \uparrow$  tone of *v. aff.* and *eff.*  $\rightarrow \downarrow$  renal blood flow but without  $\downarrow$  GFR ( $\uparrow$  FF)

higher  $\uparrow$  of sympathetic tone - during anesthesia and pain - GFR may already  $\downarrow$ 



- Regulation of renal blood flow:
- 3) Humoral Regulation
  - contribute to regulation of systemic BP and regulation of body fluids
  - NE, E (from the adrenal medulla) constriction of aff. and eff. arterioles → ↓ renal blood flow and GFR

(small impact with the exception of serious conditions, for example serious bleeding)



- Regulation of renal blood flow:
- 3) Humoral Regulation
  - contribute to regulation of systemic BP and regulation of body fluids
  - endothelin

constriction of aff. and eff. arterioles  $\rightarrow \downarrow$  renal blood flow and GFR

released locally from the impaired endothel (physiological impact - hemostasis; pathologically increased levels at the toxemia of pregnancy, acute renal failure, chronic uremia)



- Regulation of renal blood flow:
- 3) Humoral Regulation
  - contribute to regulation of systemic BP and regulation of body fluids
  - NO

continual basal production  $\rightarrow$  vasodilation  $\rightarrow$  stable renal blood flow and GFR

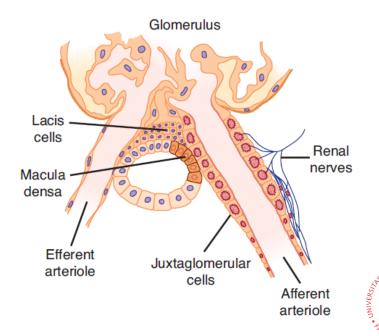
### - prostanglandins (PGE<sub>2</sub>, PGI<sub>2</sub>), bradykinin

 $\rightarrow$  vasodilation

minor impact under physiological conditions non-steroidal anti-inflammatory agents during stress!



- Regulation of renal blood flow:
  - 3) Humoral regulation
    - contribute to regulation of systemic BP and regulation of body fluids
    - Renin-angiotensine system



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