Blood-brain barrier (BBB)

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BLOOD-NERVE FIBER BARRIER – see p. A14 (2)

BBB – selective mechanism (*functional & morphological* barrier) that restricts free passage of all but **few lipid-soluble materials** from circulation to CNS parenchyma \rightarrow constant (within narrow limit) microenvironment of brain tissue.

- 1) anilino dažai i/v nudažo visus organizmo audinius, išskyrus CNS BBB darbas.
- 2) water, O₂, CO₂, gliukozė*, kai kurios amino r., lipofilinės medžiagos laisvai praeina BBB.

*highly polar molecule – enters via *facilitated diffusion* (glucose transporter GLUT 1); then other transporters (GLUT 3, GLUT 5 distribute glucose to neurons and glia)

- 3) dauguma jonų, didelės masės molekulės nepraeina pasyviai BBB reikalingas *aktyvus transportas*.
- <u>BBB makes **neuronal surface** privileged site</u> tik reikiamos medžiagos pasiekia neurono membraną, nesutrikdydamos poliarizacijos.

Pagrindinė BBB funkcija – apsaugoti neurono membranos poliarizaciją

- BBB tiksliau gal būtų vadinti **BLOOD-NEURONAL BARRIER**.

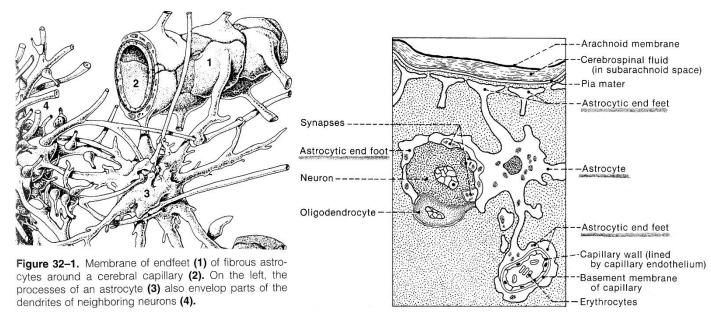
- constant extracellular ionic & molecular composition is more important for normal function of brain than for any other organ!
 - small changes in extracellular concentrations of, for example, Na⁺ or neurotransmitters (glutamate, norepinephrine) greatly alter neuronal function.
- BBB is **unidirectional**; movement of substances from brain to blood is almost unrestricted (due to bulk flow of CSF to venous system).
- BBB is *not absolute* in circumventricular organs (neurohypophysis, area postrema, etc) BBB is less effective. see p. A118 (2-3)

ANATOMICAL STRUCTURES OF BBB

- 1. Dauguma šaltinių teigia *BBB resides in vascular endothelium* unique endothelial cells (in neural capillary):
 - 1) **overlapping** (nonfenestrated)
 - 2) very **tight junctions**
 - 3) very few **pinocytotic vesicles** (little bulk flow)
 - panašūs kapiliarai randami retina, iris, inner ear, endoneurium.

Kitos galimai svarbios struktūros:

- 2. Basement membrane of capillary.
- 3. Astrocytic end feet form *continuous covering* (although 20 nm gaps may be present) of external capillary wall; taip pat glaudžiai supa neuronų kūnus (ypač izoliuoja sinapses), smegenų paviršių; astrocytic cytoplasm forms functional "extracellular" space in CNS.



- 4. Perineuronal satellite cells (astrocytes, microglia, oligodendrocytes).
- 5. Surface covering of neuronal membrane (glycoprotein, sialic acid, etc) final censor!

BLOOD-CSF BARRIER – resides in choroidal epithelium (tight junctions); features similar to BBB.

UNDER PATHOLOGIC CONDITIONS

- **BBB breaks down** → sequelae:
 - 1) *neuronal intoxication* (neuronal discharge inhibited or excited).
 - 2) vasogenic cerebral edema (plasma leakage directly into CNS tissue).
 - 3) IV contrast nudažo audinį.
 - 4) **drugs** may reach CNS tissue (e.g. antibiotics, neuroprotective drugs) "window of opportunity".

N.B. when BBB is intact, drug ability to penetrate into brain tissue depends on its lipid solubility (or presence of carrier proteins).

- pathologic conditions that break down BBB: inflammation, trauma.
 - neovessels of *brain tumors* have no BBB (capillaries are fenestrated, without tight junctions, not surrounded by astrocytic feet).

BIBLIOGRAPHY for ch. "General Histology, Myelination, BBB" → follow this LINK >>