A POROELASTIC APPROACH TO THE HYDROCEPHALUS MODELING

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Cerebrospinal fluid

Cerebrospinal fluid (CSF, liquor) — is the biological liquid around the brain and spinal cord. In the brain cerebrospinal fluid is concentrated in the four ventricles located in the center of the skull.

The total volume of cerebrospinal fluid is approximately 150 ml and it is produced at a rate of 450 ml per day (thus replacing itself three times a day).

CSF functions:

- CSF removes waste products from cerebral metabolism and supplies nutrients to nervous system tissue;
- CSF protects the brain and spinal cord from trauma;
- CSF is necessary to maintain "constant intracranial pressure"



Images are generated by Life Science Databases(LSDB). - from Anatomography website maintained by Life Science Databases(LSDB)

Cerebrospinal fluid circulation



Hydrocephalus

It is important to maintain a continuous equilibrium between the secretion, circulation and resorption of the CSF. In the case of imbalance, the fluid accumulates in the system causing a condition called hydrocephalus



Disruption of liquor production or its movement

Disruption of absorption into the circulatory system





Hydrocephalus

The Monro Kellie Doctrine

The volume of the skull is a constant which contains:

- Brain tissue (parenchyma) –80 85% intracranial space,
- Blood 6 8% intracranial space,
- CSF 5 15% intracranial space.

An increase in the volume of one component is compensated for by a decrease in the volume of one or more of the other components



Background on mathematical modeling of intracranial dynamics

- ➤<u>Hakim et al., (1971, 1976)</u> the first authors to use the methods of continuum mechanics to study hydrocephalus. They suggested that the brain parenchyma can be seen as an "open-cell sponge made of viscoelastic material."
- <u>Nagashima, Tamaki et al., (1986, 1987)</u> formalized this notion of <u>Hakim et al.</u> in terms of poroelasticity theory and the finite element method.
- ➢ Pena et al. (1999) improved Nagashima's model. The time-dependent evolution of ventricular expansion was presented in the model.
- ➢ Most studies using poroelasticity theory are based on simplified geometries, where the brain is assumed to be either spherical or cylindrical: (Kaczmarek, et al., 1997), (G Tenti et al., 1999), (Levine, 1999, 2000, 2008), (Smillie et al., 2005), (Sobey & Wirth 2006), (Tully & Ventikos 2009).
- Extension of poroelasticity models to include detailed mechanisms of transport between cerebral blood and extracellular fluid / CSF: (Tully & Ventikos 2011), (Vardakis et al., 2013), (Guo, Vardakis et al., 2018).

Mathematical modeling of hydrocephalus*



4 networks :

- Arterial blood (a)
- Arteriole/capillary blood (c)
- Venous blood (v)

• CSF (e)

The fluid flow direction. For example, from the arterial network to the capillary.

It is assumed that direct fluid transfer from the arterial network to the CSF or venous network does not occur.

^{*} Tully, Ventikos: Cerebral water transport using multiple-network poroelastic theory: application to normal pressure hydrocephalus // J. Fluid Mech. (2011), 667

Final system

 $\mu \Delta \boldsymbol{u} + (\mu + \lambda) \nabla (div \, \boldsymbol{u}) - (\alpha_a \nabla p^a + \alpha_c \nabla p^c + \alpha_e \nabla p^e + \alpha_v \nabla p^v) = 0 \tag{1}$

$$-\frac{k^a}{\mu^a}\,\Delta p^a + |\dot{s}_{a\to c}| = 0\tag{2}$$

$$-\frac{k^{c}}{\mu^{c}} \Delta p^{c} - |\dot{s}_{a \to c}| + |\dot{s}_{c \to e}| + |\dot{s}_{c \to v}| = 0,$$
(3)

$$-\frac{k^{e}}{\mu^{e}} \Delta p^{e} - |\dot{s}_{c \to e}| + |\dot{s}_{e \to v}| = 0,$$
(4)

$$-\frac{k^{\nu}}{\mu^{\nu}}\,\Delta p^{\nu}-|\dot{s}_{c\to\nu}|-|\dot{s}_{e\to\nu}|=0$$

 \boldsymbol{u} – brain tissue displacement, p – pore pressure fluid

(1) - equilibrium equation
(2) - (5) - mass conservation law for pore fluids

 $+|\dot{s}_{x\to y}| - \text{fluid out-flux from network } x \text{ to network } y;$ $-|\dot{s}_{y\to x}| - \text{fluid in-flux into the network } y \text{ from the network } x.$ The transfer is considered to be driven by a hydrostatic pressure gradient:

$$\dot{s}_{y \to x} = -\gamma_{yx} \left[p^x - p^y \right]$$



Boundary conditions

At the ventricle wall Γ_V :

• The stress must be continuous across the wall:

$$2 \,\mu \varepsilon(\boldsymbol{u}) \cdot \boldsymbol{n} + \lambda \epsilon(\boldsymbol{u}) \boldsymbol{n} = \sum_{i=a,c,e,v} (\alpha^i - 1) p^i \,\boldsymbol{n}$$

• The production of CSF from the blood results in a pressure drop in the arteriole/capillary blood network:

$$-k_{c \to ventricle} \nabla p^c \boldsymbol{n} = -Q_p$$



- $k_{c \rightarrow ventricle}$ is the resistance of the flow from the capillary network to the ventricles (through the choroid plexus).
- There is no flow into or out of the arterial and venous blood networks:

$$\nabla p^a \boldsymbol{n} = \nabla p^v \boldsymbol{n} = 0$$

• The mass of fluid in the ventricles is conserved (with d being the diameter of the cerebral aqueduct):

$$Q_p = \frac{\pi d^4}{128 \,\mu L} \left(p^e |_{\Gamma_V} - p^e |_{\Gamma_S} \right) - \oint_{Ventricle} \left(-\frac{k^e}{\mu^e} \nabla p^e \right) \cdot \mathbf{n} dS$$

Boundary conditions

At the skull Γ_S :

• The displacement is zero:

 $\boldsymbol{u}=0$

• There is no flow into or out of the capillary blood network at the skull :

$$\nabla p^c \boldsymbol{n} = 0$$

• The blood pressures are given by physically relevant arterial and venous blood pressures:

$$p^a = p_{arterial}, p^v = p_{venous},$$

• The absorption of CSF into the venous network results in a pressure rise:

$$p^e = p^v + \mu_e R Q_0$$

 μ_e –is the viscosity of the CSF,

- R is the resistance to outflow through the arachnoid granulations,
- Q_0 is the out-flux of CSF at the skull





Parameter space

The range of γ_{ac} , γ_{cv} , γ_{ce} , γ_{ev} : $10^{-4} \frac{D}{Ns}$ to $10^4 \frac{D}{Ns}$.

Each of the parameters γ_{ac} , γ_{cv} , γ_{ce} , γ_{ev} independently took values from a fifteen-element set covering the range of parameters

$$\alpha_a = \alpha_c = \alpha_v = \alpha_e = 0,99.$$

The calculation was carried out by the explicit Runge-Kutta method with automatic selection of the integration step and method order

Results: ventricular wall displacement



Fig.: Ventricular wall displacement dependence on γ_{ac} , γ_{cv} , γ_{ce} with $\gamma_{ev} = 100 \frac{D}{Ns}$

- Decrease γ_{ce} diminishing of the mutual influence of the capillary-cerebrospinal fluid component \rightarrow transfer pulsation diminishing \rightarrow CSF retention in the ventricles \rightarrow increased ventricular wall displacement
- Increase γ_{ac} increased the mutual influence of the arterial capillary component \rightarrow correct distribution of the pulse wave from arterial to capillary component \rightarrow small ventricular wall displacement
- Decrease γ_{ac} arterial-capillary flow obstruction \rightarrow significant displacement of the ventricular wall and decrease capillary pressure
- Increase γ_{cv} increased the mutual influence capillary-venous component \rightarrow overload of the venous component and inverse effect on the cerebrospinal fluid component \rightarrow increased ventricular wall displacement

Results: The pressure on the ventricular wall



- Decrease γ_{ce} capillary flow obstruction \rightarrow capillary pressure increase
- Decrease γ_{ac} arterial-capillary flow obstruction \rightarrow capillary pressure increase
- Increase γ_{cv} corresponds to increased capillary outflow \rightarrow capillary pressure decrease

Fig.: Capillary pressure dependence on γ_{ac} , γ_{cv} , γ_{ce} with $\gamma_{ev} = 100 \frac{D}{Ns}$

Results: The range of physiological norm parameters



Fig.: The range of physiological norm parameters in the parameter space γ_{ce} , γ_{cv} , γ_{ac}

In accordance with the physiological norm, the walls of the ventricles are displaced by no more than 2 mm. The range of physiological norm for capillary pressure is from 15 to 30 mmHg.

Results: Numerical simulation

The governing poroelastic equations were solved with the finite element open package FreeFem++. For finite element modeling we used triangular elements with second order approximation.





Fig.: Brain tissue displacement [mm] in $\gamma_{ce} = 2.1 * 10^3 \frac{D}{N s}$ $\gamma_{cv} = 1.06 * 10^3 \frac{D}{N s}$ $\gamma_{ac} = 1.06 * 10^3 \frac{D}{N s}$ $\gamma_{ev} = 4.5 * 10^3 \frac{D}{N s}$ (range of physiological norm parameters)

Fig.: Brain tissue displacement [mm] in

$$\gamma_{ce} = 1 * 10^4 \frac{D}{Ns}$$
 $\gamma_{cv} = 1 * 10^4 \frac{D}{Ns}$
 $\gamma_{ac} = 1.06 * 10^4 \frac{D}{Ns}$ $\gamma_{ev} = 4.5 * 10^4 \frac{D}{Ns}$

Results: Numerical simulation



Fig.: Brain tissue displacement in

$$\gamma_{ce} = 1.0 * 10^{-2} \frac{D}{N s}$$
, $\gamma_{cv} = 1.0 * 10^{-2} \frac{D}{N s}$, $\gamma_{ac} = 1.06 * 10^{-2} \frac{D}{N s}$, $\gamma_{ev} = 4.5 * 10^{-2} \frac{D}{N s}$

Conclusion

- 1. The model allows us to describe both a healthy state of an organism and a state of an organism at hydrocephalus
- 2. The model behavior correlates with the actual mechanisms of blood and CSF circulations
- 3. Calculations for simple geometry can be used to estimate the behavior of quantities in complex geometry