

# AI-supported Modelling of Brain tissue as Soft Multiscale Multiphysics (Poroelastic) medium

January 21, 2022

- Ph.D. in Structural and Materials engineering - Polytechnique University of Madrid.

# Introduction

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- Objective: simulation of poroelastic media such as **Brain tissue** (providing a digital twin).

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- A **deformable** porous solid structure (elastic, hyperelastic, viscoelastic etc.) (dry sponge)

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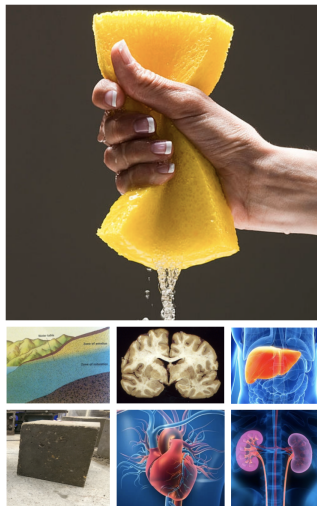


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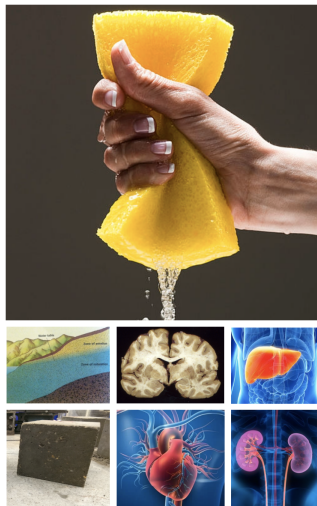


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- Rate/history dependent
- Sponge deformation causes fluid flow/percolation and vice versa

# The problem

- Brain is a soft poroelastic/poro-inelastic tissue.

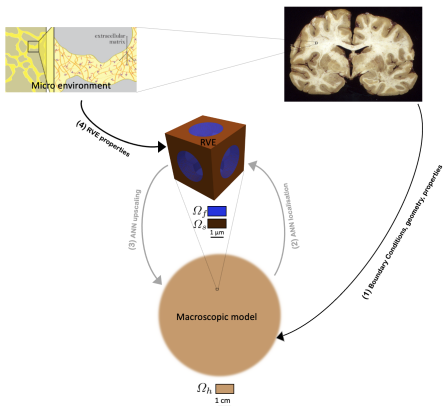


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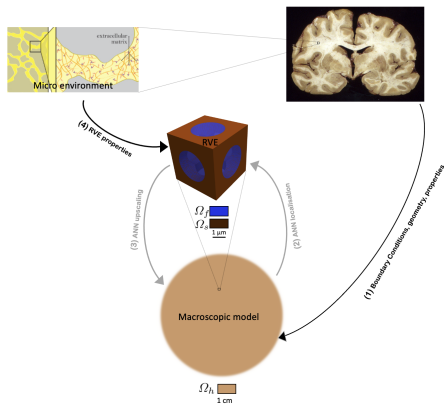


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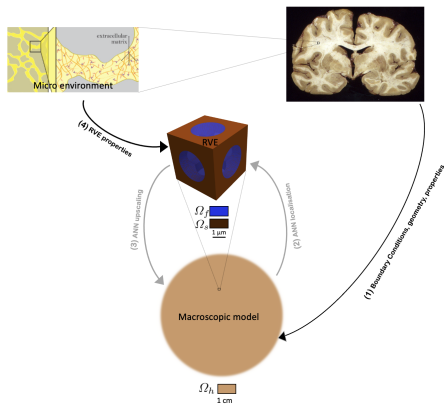


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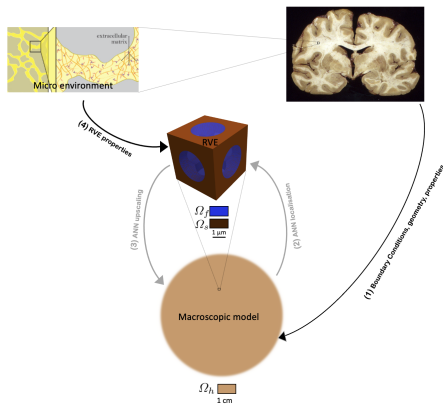


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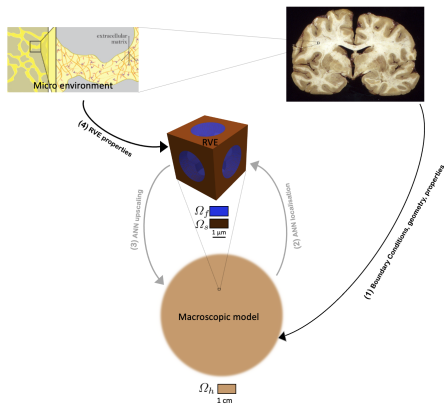


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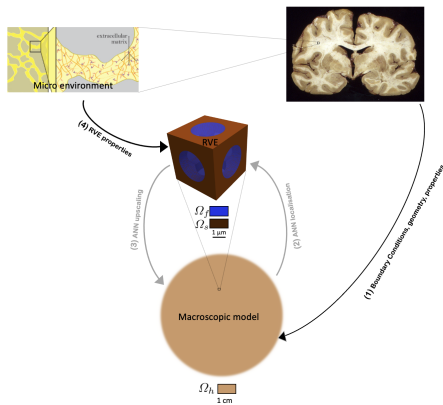


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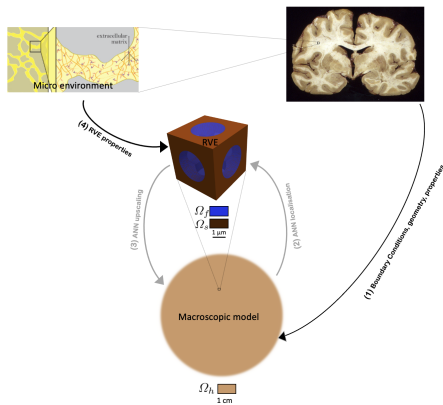


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- Paths (4) is to reach appropriate RVE based on microscopic images.



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- Are we Brain Washed during sleep?
- Such (mostly) microscopic changes are not obvious at macroscopic level while they have significant effects in material response.
- Thus: considering solid deformation is crucial even if we are only interested in fluid diffusion for drug delivery :)

remodelling based

$$\begin{aligned}
 0 = & \int_{\partial \mathcal{B}} \Delta \mathbf{t}_s \cdot \delta \mathbf{u}^{(0)} \, dS \\
 & - \int_{\mathcal{B}} \left( \tilde{\mathbb{C}} : \Delta \boldsymbol{\varepsilon}^{(0)} - \tilde{\boldsymbol{\alpha}} \Delta p^{(0)} \right) : \nabla_{\mathbf{x}} \delta \mathbf{u}^{(0)} \, dV \\
 & + \int_{\mathcal{B}} \left( \frac{1}{M} \Delta \dot{p}^{(0)} \right) \delta p^{(0)} \, dV \\
 & + \int_{\partial \mathcal{B}} (\mathbf{K} \Delta \nabla_{\mathbf{x}} p^{(0)}) \cdot \mathbf{n} \delta p^{(0)} \, dS \\
 & - \int_{\mathcal{B}} (\mathbf{K} \Delta \nabla_{\mathbf{x}} p^{(0)}) \cdot \nabla_{\mathbf{x}} \delta p^{(0)} \, dV \\
 & + \int_{\mathcal{B}} (\tilde{\boldsymbol{\alpha}} : \Delta \boldsymbol{\varepsilon}) \delta p^{(0)} \, dV \\
 & \forall \delta \mathbf{u}^{(0)}, \delta p^{(0)}.
 \end{aligned}$$

where

$$(\tilde{\mathbb{C}}, M, \tilde{\boldsymbol{\alpha}}, \mathbf{K}) = \text{ANN}(\text{microscale properties}(\text{passive effects}, \text{active effects}))$$

pros: non-formulated/structured passive and active changes, easier to develop

cons: B.C.s, complex to use

$$\begin{aligned}
 \nabla_{\mathbf{y}}^2 \mathbf{W}^T - \nabla_{\mathbf{y}} \mathbf{P} + \mathbf{I} &= \mathbf{0} \quad \text{in } \Omega_f \\
 \nabla_{\mathbf{y}} \cdot \mathbf{W}^T &= \mathbf{0} \quad \text{in } \Omega_f \\
 \mathbf{W} &= \mathbf{0} \quad \text{on } \Gamma \\
 \langle \mathbf{P} \rangle_f &= \mathbf{0}
 \end{aligned}$$

with 7 solid problems

$$\begin{aligned}
 \nabla_{\mathbf{y}} \cdot (\mathbb{C} \xi_{\mathbf{y}}(\mathcal{A})) &= \mathbf{0} \quad \text{in } \Omega_s \\
 (\mathbb{C} \xi_{\mathbf{y}}(\mathcal{A})) \mathbf{n} + \mathbb{C} \mathbf{n} &= \mathbf{0} \quad \text{on } \Gamma \\
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ALE based

$$\begin{aligned}
 0 = & + \int_{\Omega_h} \Delta \mathbf{P}_E : \nabla_{\mathbf{X}} \delta \mathbf{u}^{(0)} dV - \int_{\partial \Omega_h} \Delta \mathbf{t} \cdot \delta \mathbf{u}^{(0)} dS \\
 & - \int_{\Omega_h} \Delta (\hat{\mathbf{G}}^{(0)} \mathbf{K} (\hat{\mathbf{F}}^{(0)})^{-T} \nabla_{\mathbf{X}} \rho^{(0)}) \cdot \nabla_{\mathbf{X}} \delta \rho^{(0)} dV \\
 & + \int_{\partial \Omega_h} \Delta (\hat{\mathbf{G}}^{(0)} \mathbf{K} (\hat{\mathbf{F}}^{(0)})^{-T} \nabla_{\mathbf{X}} \rho^{(0)}) \cdot \mathbf{N}_e \delta \rho^{(0)} dS \\
 & + \int_{\Omega_h} \Delta \langle \mathbf{G}^{(0)T} : \nabla_{\mathbf{Y}} \dot{\mathbf{u}}^{(1)} \rangle_s \delta \rho^{(0)} dV \\
 & - \int_{\Omega_h} \Delta (\hat{\mathbf{G}}^{(0)T} : \nabla_{\mathbf{X}} \dot{\mathbf{u}}^{(0)}) \delta \rho^{(0)} dV
 \end{aligned}$$

where

$$\begin{aligned}
 \hat{\mathbf{F}}^{(0)} &= \langle \nabla_{\mathbf{X}} \mathbf{u}^{(0)} + \nabla_{\mathbf{Y}} \mathbf{u}^{(1)} \rangle_s + \mathbf{I} \\
 \hat{\mathbf{G}}^{(0)} &= \det \hat{\mathbf{F}}^{(0)} (\hat{\mathbf{F}}^{(0)})^{-1} \\
 \nabla_{\mathbf{Y}} \mathbf{u}^{(1)} &= ANN \left( \nabla_{\mathbf{X}} \mathbf{u}^{(0)}, \rho^{(0)} \right)
 \end{aligned}$$

$$\mathbf{0} = \nabla_{\mathbf{Y}} \cdot \mathbf{P}^{(0)} \quad \text{in } \Omega_s$$

$$\mathbf{0} = \left( \mathbf{P}^{(0)} + \rho^{(0)} \mathbf{G}^{(0)T} \right) \cdot \mathbf{N}, \quad \text{on } \Gamma$$

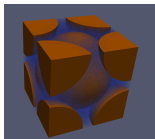
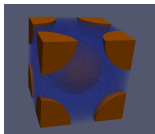
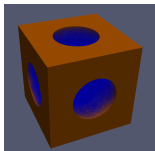
where

$$\mathbf{P}^{(0)} = \frac{\partial \Psi^{(0)}}{\partial \mathbf{F}^{(0)}}$$

pros: compatible with energy-based techniques (Hyperelasticity, phase-field fracture/cut, growth etc.), more accurate, more stable numerically, B.C. etc.

cons: it was very difficult to develop and formulate :)

# Why starting from microstructure?



# Energy distribution

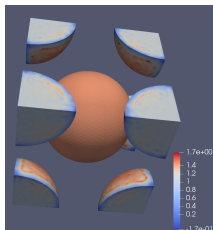


Figure 3

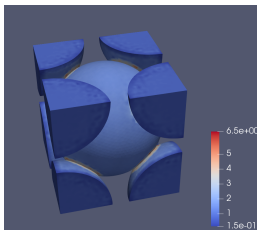


Figure 4

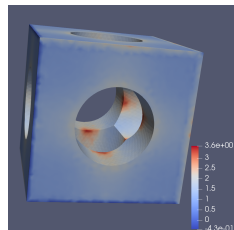


Figure 5

The RVE should reasonably match the tissue's ultrastructure.

# Some results

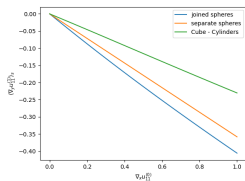


Figure 6

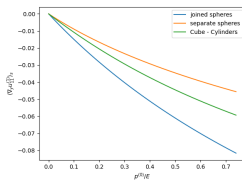


Figure 8

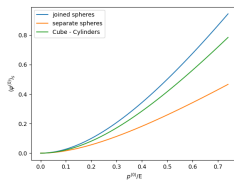


Figure 10

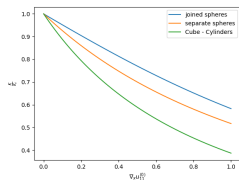


Figure 7

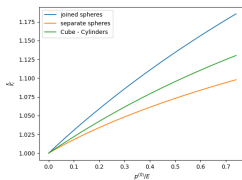


Figure 9

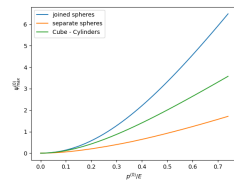
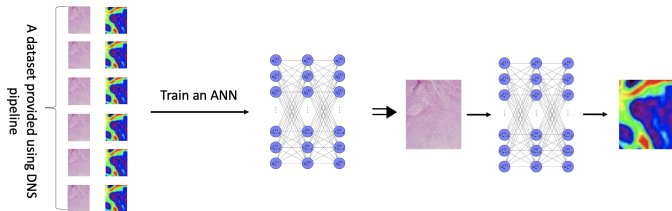
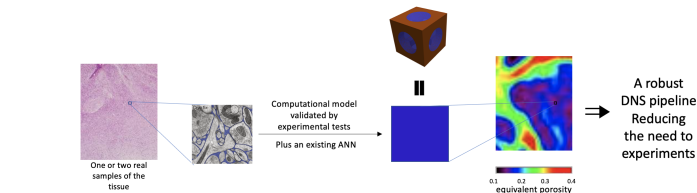


Figure 11



# Potential work package



# Concluding remarks

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- The importance and considerable influence of RVE geometry is shown in terms of both average quantities and energy concentration.
- Using some experimental results from the literature, together with our simulations we highlighted the importance of considering solid deformation and FSI features to reach a realistic prediction of fluid diffusion in brain tissue.

Thank you for your attention



Anna Golebiewska and et al.

Patient-derived organoids and orthotopic xenografts of primary and recurrent gliomas represent relevant patient avatars for precision oncology.

*Acta Neuropathologica*, 140(6):919–949, 2020.



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