## Multiscale Modelling of 3-Dimensional Brain Tissue with Capillary Distribution

Abbas Shabudin College of Engineering Universiti Malaysia Pahang Gambang, Pahang abbasshabudin@gmail.com

Stephen Payne Institute of Biomedical Engineering University of Oxford Oxford, UK stephen.payne@keble.ox.ac.uk Mohd Jamil Mohamed Mokhtarudin College of Engineering Universiti Malaysia Pahang Gambang, Pahang mohdjamil@ump.edu.my

Wan Naimah Wan Ab Naim Faculty of Mechanical and Automotive Engineering Technology Universiti Malaysia Pahang Pekan, Pahang wannaimah@ump.edu.my Wahbi K.El-Bouri Institute of Biomedical Engineering University of Oxford Oxford, UK wahbi.el-bouri@eng.ox.ac.uk

Nik Abdullah Nik Mohamed Faculty of Mechanical and Automotive Engineering Technology Universiti Malaysia Pahang Pekan, Pahang nikabdullah @ump.edu.my

## ABSTRACT

Existing brain model has been developed to study brain oedema formation in ischaemic stroke, assumed the brain as a homogenized structure and ignored the effects of blood capillary of the brain. Thus, the aim of this study is to reconsider the effects of capillary in the brain model through multiscale approach using asymptotic expansion homogenization (AEH) technique. AEH is applied to the existing governing equations of the brain, resulting in new governing equations consist of 6 homogenized macroscale equations and 4 microscale cell problems. Actual brain capillary geometry is developed based on actual capillary network distribution data generated using modified spanning tree method. The microscale cell problems are then solved in actual brain capillary geometry in order to obtain four important parameters, namely the hydraulic conductivity, homogenous Biot's coefficient and elastic stiffness tensor. From the result, the distribution matrix obtained for hydraulic conductivity is not isotropic. This problem can be improved by increasing the volume of the actual capillary geometry. For homogenous Biot's coefficient, the matrix obtained is isotropic, however the reliability of the result obtained can be improved by solving the cell problem in multiple capillary geometries. For elastic stiffness tensor, it can be concluded that this parameter does not significantly affected the macroscale equations of bigger brain. All these parameters are required later in order to solve the homogenized macroscale equations for the investigation of brain diseases such as ischaemic stroke and dementia.

*KEYWORDS:* actual capillary geometry, ischaemic stroke, asymptotic expansion homogenization, macroscale equations, microscale cell problems

## ACKNOWLEDGMENT

The research is supported by Fundamental Research Grant Scheme FRGS/1/2018/TK03/UMP/02/15 (RDU 190132) and PGRS1903153. Wan Naimah Wan Ab Naim is the recipient of Universiti Malaysia Pahang (UMP) Post-Doctoral Fellowship in Research.

## REFERENCES

- M. Katan and A. Luft A, "Global burden of stroke,"Seminars in neurology, vol. 38 (2), pp. 208-211), 2018.
- [2] B. C. Campbell, D. A. De Silva, M. R. Macleod, S. B. Coutts, L. H. Schwamm, S. M. Davis, and G. A. Donnan, "Ischaemic stroke," Nature Reviews Disease Primers, vol. 5(1), pp. 1-22, 2019.
- [3] E. S. Donkor, "Stroke in the 21st century: A snapshot of the burden, epidemiology, and quality of life," Stroke research and treatment, 2018.
- [4] K. W. Loo and S. H. Gan, "Burden of stroke in Malaysia,". International Journal of Stroke, vol. 7(2), pp. 165-167, 2012.
- [5] S. Vidale, and E. C. Agostoni, "Organizing Healthcare for Optimal Acute Ischemic Stroke Treatment," Journal of Clinical Neurology. Vol. 16(2), pp. 183, 2020.
- [6] S. Michinaga and Y. Koyama, "Pathogenesis of brain edema and investigation into anti-edema drugs," International journal of molecular sciences, vol. 16(5), pp. 9949-9975, 2015.
- [7] J. Bardutzky and S. Schwab, "Antiedema therapy in ischemic stroke," Stroke, vol. 38(11), pp. 3084-3094, 2007.
- [8] R. Penta and D. Ambrosi, "The role of the microvascular tortuosity in tumor transport phenomena," Journal of theoretical biology, vol. 364, pp. 80-97, 2015.
- [9] R. Penta and J. Merodio, "Homogenized modeling for vascularized poroelastic materials," Meccanica, 52(14), pp. 3321-3343, 2017.
- [10] X. Fan and H. Markram, "A brief history of simulation neuroscience," Frontiers in Neuroinformatics, vol. 13, pp. 32, 2019.
- [11] M. Peyrounette, Y. Davit, M. Quintard, and S. Lorthois,"Multiscale modelling of blood flow in cerebral microcirculation: Details at capillary scale control accuracy at the level of the cortex," PloS One, vol. 13(1), pp. e0189474, 2018.
- [12] M. J. M. Mokhtarudin and S. J. Payne, "Mathematical model of the effect of ischemia-reperfusion on brain capillary collapse and tissue swelling," Mathematical Biosciences, vol. 263, pp. 111-120, 2015.
- [13] W. K. El-Bouri and Payne, "Multi-scale homogenization of blood flow in 3-dimensional human cerebral microvascular networks," Journal of theoretical biology, vol. 380, pp. 40-47, 2015
- [14] F. Cassot, F. Lauwers, C. Fouard, S. Prohaska, and V. Lauwers-Cances, "A novel three-dimensional computer-assisted method for a quantitative study of microvascular networks of the human cerebral cortex," Microcirculation, vol. 13(1), pp. 1-18, 2006.
- [15] A. Shabudin, M. J. M. Mokhtarudin, S. Payne, and N. A. N. Mohamed, "Application of asymptotic expansion homogenization for vascularized poroelastic brain tissue," 2018 IEEE-EMBS Conference on Biomedical Engineering and Sciences (IECBES), pp. 413-418, 2018.
- [16] A. Shabudin, M. J. M. Mokhtarudin, S. Payne, and N. A. N. Mohamed, "Investigation of the capillary effects toward brain tissue poroelastic properties using asymptotic expansion homogenization," 6<sup>th</sup> International Conference on Computational and Mathematical Biomedical Engineering-CMBE2019, 2019.
- [17] M. A. Biot, "General theory of three-dimensional consolidation," Journal of applied physics, 12(2), pp. 155-164, 1941.