Subject-specific Modeling of Intra-arterial Nanoparticle Delivery for Glioma Treatments

Shaolie S. Hossain*'[§] and Shailendra Joshi[†]

^{*}Department of Molecular Cardiology, Texas Heart Institute, 6770 Bertner Avenue, Houston, TX 77030, USA e-mail: shossain@texasheart.org [§]Institute for Computational Engineering and Sciences, The University of Texas at Austin, 201 E. 24th St., Austin, TX 78712, USA

> [†]College of Physicians and Surgeons, Columbia University, 161 Fort Washington Avenue, New York, NY 10032, USA email: sj121@columbia.edu

ABSTRACT

Nanoparticles are considered important vehicles for targeted release of drugs to treat brain tumors or gliomas, and other focal neurological diseases. The regional deposition of surface-functionalized nanoparticle is a dynamic balance between the forces of particle attachment and hydrodynamic forces that tend to dislodge them [1]. Few computational models have explored this relationship of regional nanoparticle delivery after intra-arterial (IA) injections in the Circle of Willis. In this work we used computational tools to investigate the liposomal deposition under condition of cerebral hypoperfusion for different particle characteristics. The results were compared with observations from *in vivo* rodent experiments conducted with optically tagged nanoparticles, which can provide important insight into the pharmacokinetics of IA drug delivery [2]. Both experimental and simulation data supported the improvement in regional tissue deposition with transient cerebral hypoperfusion, cationic charge and larger particle size. Our model, which incorporates nanoparticle design parameters, intra-arterial injection profile, subject-specific anatomical features and realistic regional blood flow, provides the conceptual framework to understand and improve intra-arterial drug delivery to potentially avoid complications and failures observed during IA chemotherapy in glioma treatments [3].

REFERENCES

- S.S. Hossain, Y. Zhang, X. Liang, F. Hussain, M. Ferrari, T.J.R. Hughes and P. Decuzzi, "in silico Vascular Modeling for Personalized Nanoparticle Delivery," Nanomedicine, 8 (2013), 343-357. DOI: 10.2217/NNM.12.124
- [2] S. Joshi, J.R. Cooke, D.K. Chan, J.A. Ellis, S.S. Hossain, R.P. Singh-Moon, et al. "Liposome size and charge optimization for intraarterial delivery to gliomas," *Drug Delivery and Translational Research*. (2016). DOI: 10.1007/s13346-016-0294-y
- [3] J.A. Ellis, M. Banu, S.S. Hossain, R. Singh-Moon, S.D. Lavine, J.N. Bruce and S. Joshi, "Reassessing the Role of Intra-Arterial Drug Delivery for Glioblastoma Multiforme Treatment," *Journal of Drug Delivery* 2015 (2015) 15. DOI:10.1155/2015/405735