Nanoparticles transport in Glioblastoma from intracranially-administered thermosensitive hydrogels

Giulia Brachi^{1,2}, Gianluca Ciardelli², Robert Rostomily³, Andrei Mikheev³, Clara Mattu^{1,2}, Mauro Ferrari^{1,4}

- 1. Department of Nanomedicine, Houston Methodist Research Institute, Houston, TX, USA
- 2. Department of Mechanical and Aerospace Engineering, Politecnico di Torino, Torino, ITALY
- 3. Center for Neuroregeneration, Houston Methodist Research Institute, Houston, TX, USA
- 4. Department of Medicine, Weill Cornell Medical College, New York, NY, USA

Glioblastoma multiforme (GBM) is the most common and aggressive primary brain tumor in adults¹. Intracranial (IC) drug delivery is a promising strategy to treat GBM, because of the potential to bypass the blood-brain barrier (BBB), reduce systemic side effects and enhance drug concentration at the tumor site^{2,3}.

However, high interstitial fluid pressure in GBM results in rapid elimination of IC-administered treatments from the tumor bulk, thus requiring new strategies to increment treatment retention. In this work, thermosensitive hydrogels loaded with multifunctional polymer nanoparticles (HG-NPs) were designed and characterized for IC drug delivery in GBM.

HG-NPs and free NPs were IC administered in tumor free mice and in a highly infiltrative GBM model⁴ (18 days after tumor inoculation) and their transport kinetics were investigated using complementary 2D/3D In vivo imaging (IVIS) system and ex vivo fluorescence imaging.

HG-NPs resulted in reduced treatment clearance after injection, high tissue penetration ability, enhanced tumor coverage and significant increase in long-term retention inside the brain, thus warranting further investigation as novel approach for GBM treatment.

References:

- (1) Nam, J.Y.; J. Oncol. Pract. 2017, 13 (10): 629–638.
- (2) Sarkaria, J. N.; Neuro. Oncol. 2018, 20 (2):184-191.
- (3) Bastiancich, C.; *J Control Release*. 2016, 225:283-93.
- (4) Wakimoto, H.; Cancer Res. 2010, 69 (8): 3472–3481.