

## **Early changes in tumor perfusion from dynamic contrast enhanced T1-weighted MRI following neural stem cell-mediated therapy of recurrent high grade glioma correlates with overall survival**

### **SUPPLEMENTAL MATERIALS AND METHODS**

#### **MRI Scanning protocol**

All patients underwent the same DCE-MRI scanning protocol as follows: repetition time/echo time (TR/TE):9.3/4.29ms; flip angle:15°; acquisition matrix  $81 \times 192$ ; field of view (FOV)  $192 \times 280$  (mm<sup>2</sup>); slice thickness 5mm; number of slices 16; number of dynamic phase 50; temporal resolution 5s. At the fifth time point of the DCE-MRI data acquisition, 0.2 mmol/kg body weight of Gd-BOPTA (Multihance, Bracco, Italy) was administered intravenously with a power injector at a rate of 3.5 ml/s, followed by a bolus injection of a 30ml saline flush. Three different flip angle (2°, 5° and 10°) pre-contrast T1-weighted images were acquired for absolute quantification of  $T_1$  relaxation time. The total acquisition time for dynamic scan was 4min 10sec.

#### **Quantification of Perfusion Parameters**

Pre-contrast  $T_1$  was measured by using pre-contrast T1-weighted images acquired at three different flip angles [1]. Contrast agent concentration time curves were computed voxel-wise using Signal intensity time DCE-MRI data and pre-contrast  $T_1$ . Automatic arterial input function was extracted by using the algorithm described in [2]. A three-compartment pharmacokinetic leaky tracer kinetic model (LTKM) was fitted to the concentration time curve and fractional plasma volume ( $V_p$ ), permeability ( $K^{trans}$ ), leakage ( $\lambda^{tr}$ ) were estimated as previously reported [3]. An in-house JAVA-based software was used for all DCE computations.

## References

1. Liberman G, Louzoun Y, Ben Bashat D. T<sub>1</sub> Mapping using variable flip angle SPGR data with flip angle correction. *J Magn Reson Imaging*. 2014;40(1):171-180. doi:10.1002/jmri.24373.
2. Singh A, Rathore RKS, Haris M, Verma SK, Husain N, Gupta RK. Improved bolus arrival time and arterial input function estimation for tracer kinetic analysis in DCE-MRI. *J Magn Reson Imaging*. 2009;29:166-76.
3. Sahoo P, Rathore RKS, Awasthi R, et al. Subcompartmentalization of extracellular extravascular space (EES) into permeability and leaky space with local arterial input function (AIF) results in improved discrimination between high- and low-grade glioma using dynamic contrast-enhanced (DCE) MRI. *J Magn Reson Imaging*. 2013;38(3):677-688.