

### 3T multiecho gradient-recalled echo BOLD MRI with high reproducibility for serial assessment of limb muscle oxygenation

Erik Hedstrom<sup>1,2</sup>, Ashish S Patel<sup>2,3</sup>, Tobias Voigt<sup>1,4</sup>, Bijan Modarai<sup>2,3</sup>, Tobias Schaeffter<sup>1,2</sup>, Alberto Smith<sup>2,3</sup>, and Eike Nagel<sup>1,2</sup>

<sup>1</sup>Division of Imaging Sciences and Biomedical Engineering, King's College London, London, Greater London, United Kingdom, <sup>2</sup>BHF Centre of Research Excellence and NIHR Biomedical Research Centre at Guy's and St Thomas' NHS Fo, London, United Kingdom, <sup>3</sup>Academic Department of Surgery, King's College London, London, Greater London, United Kingdom, <sup>4</sup>Philips Research, Clinical Research Europe

**BACKGROUND:** Current imaging methods, including BOLD MR imaging, cannot reliably quantify muscle oxygenation in patients with limb ischaemia. The BOLD response is complex and measurements have so far been performed in relatively low-resolution EPI-readout images where it may be difficult to exclude oxygenation changes from partial volume effects. The reproducibility of BOLD imaging has been low and consequently its usefulness as a clinical test has been limited. We propose a high-resolution BOLD sequence whereby edge artefacts and vessels may be excluded from measurements, analysed using a Maximum Likelihood Estimate with Rician noise correction, where noise is read in k-space, for a precise and unbiased estimation of T2\* from magnitude images. We test the hypothesis that this gives high reproducibility at baseline, and during and after transient ischaemia.

**MATERIALS AND METHODS:** The lower limbs of eight volunteers without limb disease (median age 66±3yrs) were imaged twice for reproducibility at time intervals between 1 and 190 days at 3T with a 32-channel coil (Philips Achieva, Best, NL). During each scan transient ischaemia was induced by application of a compression cuff around the thigh inflated to suprasystolic pressure for 5mins. The multi-echo multi-shot GRE BOLD images (TR 66ms, TE<sub>1</sub> 4.6 ms, ΔTE 4.6 ms, 14 echoes, 1535 Hz/px, FA 20°, FOV 300×150mm, matrix 256×128) were acquired every 2s at baseline and during the dynamic phases of ischaemic response. Data was analysed for early (20s) and late (280s) during cuffing, and for early (20s) and late (315s) after cuff deflation, and interscan reproducibility assessed. Intrascan variability was also determined in an uncuffed population (n=12; median age 67±2yrs). Regions of interest were drawn around the anterior (ant) and lateral (lat) muscle compartments and around the gastrocnemius (gc) and the soleus (sol) muscles, masking out edge artefacts and vessels.

**RESULTS:** Image resolution was superior to standard EPI BOLD (Figure 1).

Intrascan variability for each muscle group was: ant<0.3±0.5ms, lat<0.3±0.9ms, gc<0.2±0.5ms, and sol<0.1±0.6ms (all p values between intrascan time points for each muscle group=NS), Figure 2. Interscan reproducibility for baseline was: ant<1.3±1.9ms, lat<1.7±1.2ms, gc<1.0±1.0ms, and sol<0.6±2.5ms; for minimum T2\* during ischaemia ant<0.8±0.2ms, lat<0.9±1.9ms, gc<0.7±1.0ms, and sol<2.1±1.6ms; for maximum T2\* early after cuff deflation ant: <0.2±1.0ms, lat: <3.0±0.3ms, gc: <1.0±1.1ms, and sol: <0.9±0.7ms; and for T2\* late after cuff deflation ant<0.8±1.2ms, lat<2.4±1.5ms, gc<0.9±0.3ms, and sol<0.9±0.6ms (all p=NS; Figure 3). Compared with baseline, the minimum and maximum T2\* values were -19±7% and 1±7%, respectively.

**CONCLUSIONS:** The sequence and analysis proposed shows low intrascan variability and high interscan reproducibility for measurement of T2\*. The proposed technique is therefore well suited for serial assessment of limb muscle oxygenation.

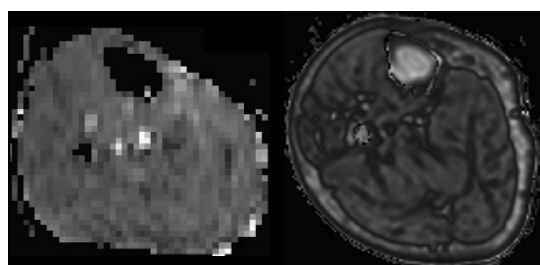


Figure 1. T2\* maps by standard EPI BOLD (left) compared with the proposed non-EPI GRE BOLD sequence (right).

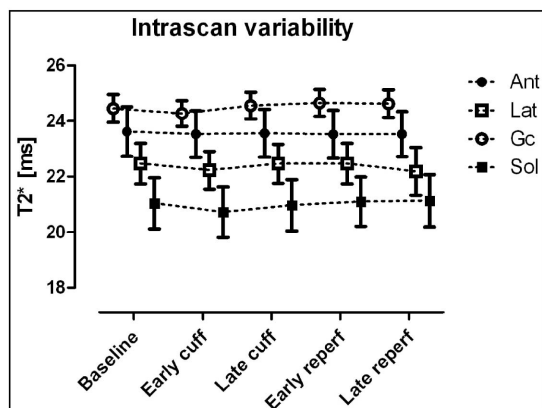


Figure 2. Intrascan variability for the four compartments in an uncuffed population. Error bars indicate standard error of the mean. Ant=anterior compartment; Lat=lateral compartment; Gc=gastrocnemius; Sol=soleus.

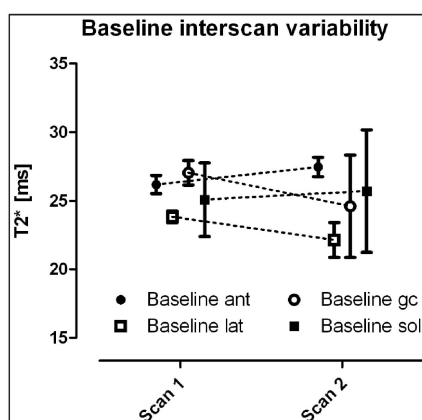


Figure 3. Interscan variability for T2\* measurements between the repeated scans in the cuffed population. The measurements for minimum, maximum, and late reperfusion T2\* values are similar, and only baseline shown for clarity. Error bars indicate standard error of the mean. Ant=anterior compartment; Lat=lateral compartment; Gc=gastrocnemius; Sol=soleus.