

## Assessment of Critical Limb Ischemia using MRI

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### Background

Critical limb ischaemia (CLI) is a manifestation of peripheral arterial disease (PAD) that describes patients with chronic ischaemic rest pain, or ischaemic skin lesions in the form of ulcers or gangrene[1]. Current treatment approaches differ, in part due to disputes over the best non-invasive method of assessing CLI. We therefore sought to determine whether MRI could provide a non-invasive objective assessment of the degree of ischemia without the use of contrast agents.

### Hypothesis/Objective

We examined the use of three MRI maps to depict vascular impairment:

$T_2^*$  images, the hypothesis being, that increased amounts of de-oxygenated blood in ischemic patients yields increased amounts of paramagnetic methemoglobin causing faster spin relaxation and a decreased  $T_2^*$  time (BOLD effect).

$T_2$  images, hypothesized to depict edemas in chronic ischemic patients. Further, the obtained  $T_2$  map may be used to correct  $T_2^*$  for water content through the use of  $T_2'$ .

$T_2'$  images, hypothesized to be superior to  $T_2^*$  for depicting methemoglobin content, since this map incorporates both relaxation due to paramagnetic effects and signal increases due to edema.

### Methods

The calfs of five patients suffering from chronic ischemia in the legs and three healthy volunteers were scanned using a Philips Intera Achieva 1.5T scanner. A fast gradient echo sequence was used to calculate  $T_2^*$  maps employing a single cross-sectional slice using eight images with echo times (TE) ranging from 6.9 ms to 83.2 ms, TR of 108.8 ms, flip angle (FA) of 30°, 2 averages, and 0.5x0.5x6 mm<sup>3</sup> voxel size. Additionally  $T_2$  maps were calculated using a turbo spin echo sequence using eight images with TE from 10 to 80 ms, TR = 108 ms, 6 averages, and 0.5x0.5x10 mm<sup>3</sup> voxel size.

Post-processing was applied to fit  $T_2$ - and  $T_2^*$  images (Figure 1) to exponential curves using the free open-source software OSIRIX[2]. Using the 2D Grow Region Confidence algorithm (multiplier: 2.5, num. of iteration: 5, initial radius: 2), the calf muscles were segmented, and the average signal intensity calculated (Figure 2). The average  $T_2$ - and  $T_2^*$  values were extracted and used to calculate the  $T_2'$  using the relation:

$$\frac{1}{T_2'} = \frac{1}{T_2} + \frac{1}{T_2^*} \Leftrightarrow T_2' = 1 / \left( \frac{1}{T_2} + \frac{1}{T_2^*} \right)$$

### Results

$T_2$  values (Figure 3A) differed significantly (P=0.011, CI95=[5.6;29.7]) between patients ( $T_2=57.4 \pm 7.9$  ms) and controls ( $T_2=39.7 \pm 3.3$  ms).  $T_2^*$  (Figure 3B) showed no significant difference (P=0.107, CI95=[-1.14;8.91]) between patients ( $T_2^*=33.8 \pm 2.6$  ms) and controls ( $T_2^*=29.9 \pm 3.1$  ms).  $T_2'$  (Figure 3C) also did not display any insignificant difference (P=0.09, CI95=[-78.56;7.17]) between patients ( $T_2'=87.6 \pm 22.7$  ms) and controls ( $T_2'=123.3 \pm 26.4$  ms).

### Discussion

Initial results did not show any significant differences between severely ischemic patients and normal controls using either  $T_2^*$  or  $T_2'$ , indicating the BOLD effect of deoxygenated blood to be too small to measure using the present sample size and settings. However,  $T_2$  values were significantly affected, hypothesized to be the result of edema caused by the severe ischemia.

### Conclusion

MRI  $T_2$  maps may prove viable for assessing severely ischemic patients quantifiably.

### References

1. Norgren, L., et al., *Inter-Society Consensus for the Management of Peripheral Arterial Disease (TASC II)*. Journal of Vascular Surgery, 2007. 45(1, Supplement 1): p. S5-S67.
2. Rosset, A., L. Spadola, and O. Ratib, *OsiriX: An Open-Source Software for Navigating in Multidimensional DICOM Images*. Journal of Digital Imaging, 2004. 17(3): p. 205-216.

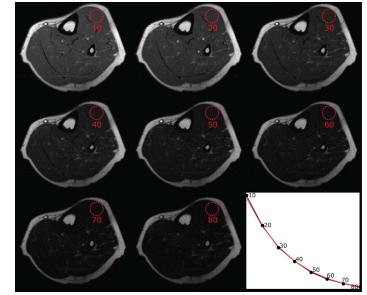


Figure 1: T2 images used to calculate T2 map.

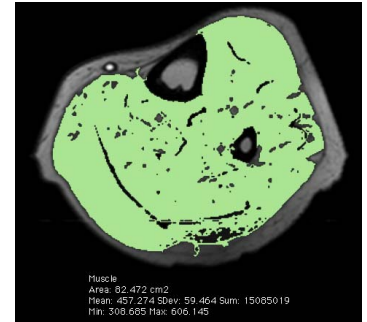


Figure 2: T2 image with segmented muscles.

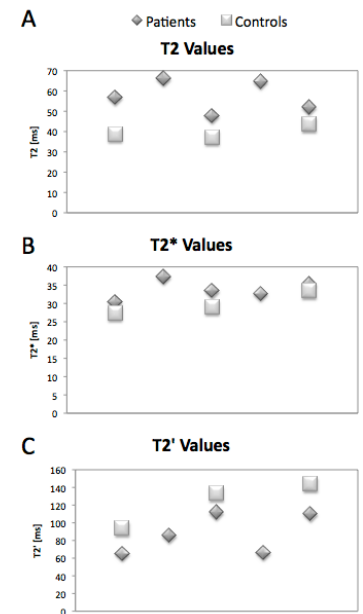


Figure 3: Results