

Scientific Paper:

Magn Reason Mater Phy (2022)

## Determination of oxygen relaxivity in oxygen nanobubbles at 3 and 7 Tesla

Emma Bluemke<sup>1</sup>, Liam A. J. Young<sup>2</sup>, Joshua Owen<sup>3</sup>, Sean Smart<sup>4</sup>, Paul Kinchesh<sup>4</sup>, Daniel P. Bulte<sup>1</sup>, Eleanor Stride<sup>1</sup>

<sup>1</sup>Department of Engineering Sciences, Institute of Biomedical Engineering, University of Oxford, Oxford, UK

<sup>2</sup>Radcliffe Department of Medicine, Oxford Centre for Clinical Magnetic Resonance Research, University of Oxford, Oxford, UK

<sup>3</sup>Clinical Center, National Institutes of Health, Bethesda, USA

<sup>4</sup>Department of Oncology, Radiobiology Research Institute, University of Oxford, Oxford, UK

## Abstract:

**Objective** Oxygen-loaded nanobubbles have shown potential for reducing tumour hypoxia and improving treatment outcomes, however, it remains difficult to noninvasively measure the changes in partial pressure of oxygen  $(PO_2)$  in vivo. The linear relationship between  $PO_2$  and longitudinal relaxation rate  $(R_1)$  has been used to noninvasively infer  $PO_2$  in vitreous and cerebrospinal fluid, and therefore, this experiment aimed to investigate whether  $R_1$  is a suitable measurement to study oxygen delivery from such oxygen carriers.

**Methods**  $T_1$  mapping was used to measure  $R_1$  in phantoms containing nanobubbles with varied  $PO_2$  to measure the relaxivity of oxygen  $(r_{10x})$  in the phantoms at 7 and 3 T. These measurements were used to estimate the limit of detection (LOD) in two experimental settings: preclinical 7 T and clinical 3 T MRI.

**Results** The  $r_{10x}$  in the nanobubble solution was 0.00057 and 0.000235 s<sup>-1</sup>/mmHg, corresponding to a LOD of 111 and 103 mmHg with 95% confidence at 7 and 3 T, respectively.

**Conclusion** This suggests that  $T_1$  mapping could provide a noninvasive method of measuring a > 100 mmHg oxygen delivery from therapeutic nanobubbles.

Keywords: quantitative MRI, MRI relaxometry, oxygen, hypoxia