## Hyperpolarized [1-<sup>13</sup>C]-acetate Renal Metabolic

## **Clearance Rate Mapping**

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Short running title: Hyperpolarized <sup>13</sup>C-acetate MRI

## SUPPLEMENTAL FIGURES.

Supplemental figure 1.

In order to verify the existence of a metabolic component in the mean transit time, 4 female rats receiving a similar furosemide treatment was processed with the proposed method. A global  $T_1$  relaxation time of 24 sec, was used to correct the <sup>13</sup>C-urea perfusion data.



**SUPPLEMENTAL FIGURE 1.** The <sup>13</sup>C-urea RBF did not differ statistically between pre ( $450 \pm 88$  ml/100 ml/min) and post administration of furosemide ( $477 \pm 96 (\pm SD)$  ml/100 ml/min; paired *t*-test: P = 0.7). Furthermore, no difference in the <sup>13</sup>C-urea MTT of  $18 \pm 5 (\pm SD)$  sec at baseline compared with  $16 \pm 4 (\pm SD)$  sec post furosemide administration (paired *t*-test: P = 0.6) was found.

Supplemental figure 2.

Whole blood was extracted from two healthy rats into sodium heparin vacuum tubes. The blood was stored at 5° C. Prior to the experiment the blood was heated and maintained at 37° C. A volume of 4.5 ml blood was mixed with hyperpolarized <sup>13</sup>C-acetate (0.5 ml) prior to placement in the scanner. The MR experiment was acquired over 120 s (120 acquisitions), with a constant flip angle of 10°. The single exponential decay was fitted in MATLAB and corrected for RF depletion.

$$\frac{1}{T_1} = \frac{1}{T_{1_{eff}}} - \frac{\ln \cos \theta}{TR}$$
(supplement eq. 1)

Where  $\theta$  is the RF flip angle and TR is the repetition time. Using the *ex vivo* T<sub>1eff</sub> for correction, results in an underestimated absolute perfusion, lower MTT and thus larger K<sub>MTT</sub>.



**SUPPLEMENTAL FIGURE 2.** <sup>13</sup>C-acetate *in vivo* hemodynamic parameters (using whole blood T<sub>1</sub> correction). Acetate perfusion (min/100 ml/mL), mean transit time (MTT) (sec), and acetate mean transit time metabolic clearance rate  $K_{MTT}$  (min<sup>-1</sup>) before and after administration of furosemide. The mean is plotted with standard errors.