**Assessing Murine Cardiac Metabolism Using a 10 mm, 13C Optimized Cryoprobe**

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**Introduction**

 Heart failure is characterized by a profound shift in energy metabolism. The normal myocardium preferentially consumes fatty acids, with only 20 to 30 % of oxidative metabolism being supplied by glucose. Overpressure of the heart, either from high blood pressure or stenosis, causes myocardial hypertrophy and a switch to carbohydrate oxidation1. Whether the switch in substrate preference is directly causal of hypertrophy, or a byproduct of it remains debated2. We have previously measured the ability of propionate to modulate carbohydrate oxidation in the murine heart using hyperpolarized (HP) pyruvate3 and in experiments directly assessing glucose oxidation4. Here we detail initial experiments using the same tracer methods in the hypertrophic heart.

**Experimental**

 Mouse hearts (healthy or failing) were excised and aorta was cannulated. The hearts were perfused in langendorff mode (with or without propionate) in an NMR magnet with a field strength of 14.1 T equipped with a 10 mm Bruker cryoprobe (AMRIS, UF, Gainesville). Shimming the magnet was accomplished using 23Na signal from the heart. After 30 minutes of perfusion, hyperpolarized [1 – 13C]pyruvic acid was injected into the heart and 13C NMR spectra was recorded with a repetition time of 4s using 15° radiofrequency pulses.

**Results and Discussion**



**Fig.1** (left) Sum of 49 13C NMR spectra recorded subsequent to the injection of HP [1 – 13C] pyruvic acid showing metabolic activity of the heart. (right) Ratio of bicarbonate to pyruvate in perfused healthy and failing (TAC) hearts in the presence and absence of propionate.

 The Oxford HyperSense instrument readily produces signal enhancements of ~20000 fold for pyruvic acid, as measured by dissolution and measurement at room temperature and 14.1 T. Injection of the HP pyruvate into the functioning heart results in its rapid metabolism by exchange reactions with lactate and alanine as well as flux through pyruvate dehydrogenase to produce bicarbonate. The 10 mm cryoprobe supported by the MagLab provides an additional 4x gain in SNR for the measurements (Figure 1, Left). Pyruvate oxidation is increased in the hypertrophic heart (Figure 1, right).

**Conclusions**

 The 10 mm cryoprobe paired with the HyperSense allow substrate selection to be measured in a straightforward manner in the perfused mouse heart.

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