Study of real time 13C metabolic fluxes using Dynamic Nuclear Polarization and Magnetic Resonance Spectroscopy

Commonly used acronym: DNP


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SCOPE OF THE METHOD

<table>
<thead>
<tr>
<th>The Method relates to</th>
<th>Animal health</th>
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<tbody>
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<td>The Method is situated in</td>
<td>Basic Research, Translational - Applied Research</td>
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<tr>
<td>Type of method</td>
<td>Other: In vitro and in vivo method</td>
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</tbody>
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DESCRIPTION

Method keywords

hyperpolarization
13C metabolic fluxes
13C-MRS
in vitro and in vivo tool

**Scientific area keywords**

- tumor metabolism
- cardiac function
- metabolic disorders
- 13C metabolism

**Method description**

Hyper Polarized (HP) MR allows to considerably increase the sensitivity (>10,000) of MR spectroscopy and spectroscopic imaging. The hyperpolarized molecule can be injected at room temperature to a cell system or an *in vivo* model and its metabolism can be followed over a few minutes. The technique has been used *in vivo* or *in vitro* to assess changes in metabolic fluxes through glycolysis, citric acid cycle, and fatty acid synthesis. HP MR studies using 13C pyruvate showed an increase in signal to noise ratio of more than 50,000 of the substrate and its metabolites (lactate and alanine), allowing for unique dynamic mapping of metabolism using spectroscopic imaging.

**Lab equipment**

The Oxford Instruments HyperSense DNP system is combined with our 11.7T preclinical Magnetic Resonance system (Bruker Biospin) for *in vivo* application or with our Bruker Ascend 600MHz NMR system for *in vitro* application. The Hypersense allows hyperpolarization of 13C-enriched substrates, direct dissolution and cooling down of the substrates and allows consecutive injection in the vascular system of the animal under study or in cell media for intro application.

**Method status**

Published in peer reviewed journal

**PROS, CONS & FUTURE POTENTIAL**

**Advantages**

The method allows the assessment of real-time metabolic conversion (i.e. of 13C-pyruvate into 13C-lactate) with high sensitivity and non-invasively (while used *in vivo*). It constitutes a key tool to address fundamental questions in the scope of
metabolism in several fields, including oncology, cardiology, metabolic disorders, etc...

**Challenges**

A major limitation of DNP is a requirement of a long relaxation time T1 for the 13C enriched substrate. The useful monitoring time is limited to 5X T1, which includes dissolution, injection, and imaging. Hence, for pyruvate, there is a maximum of 3 min of useful measuring time. Nevertheless, the technique has already shown numerous applications for detecting treatment response in animals.

**Modifications**

No modifications are planned in the near future.

**Future & Other applications**

Metabolic disorders;
Cardiac function metabolism;
Cancer metabolism.

**REFERENCES, ASSOCIATED DOCUMENTS AND OTHER INFORMATION**

**References**

