

# Real Time Molecular Imaging of TCA Cycle Metabolism *In Vivo* By Hyperpolarized 1-<sup>13</sup>C Diethyl Succinate

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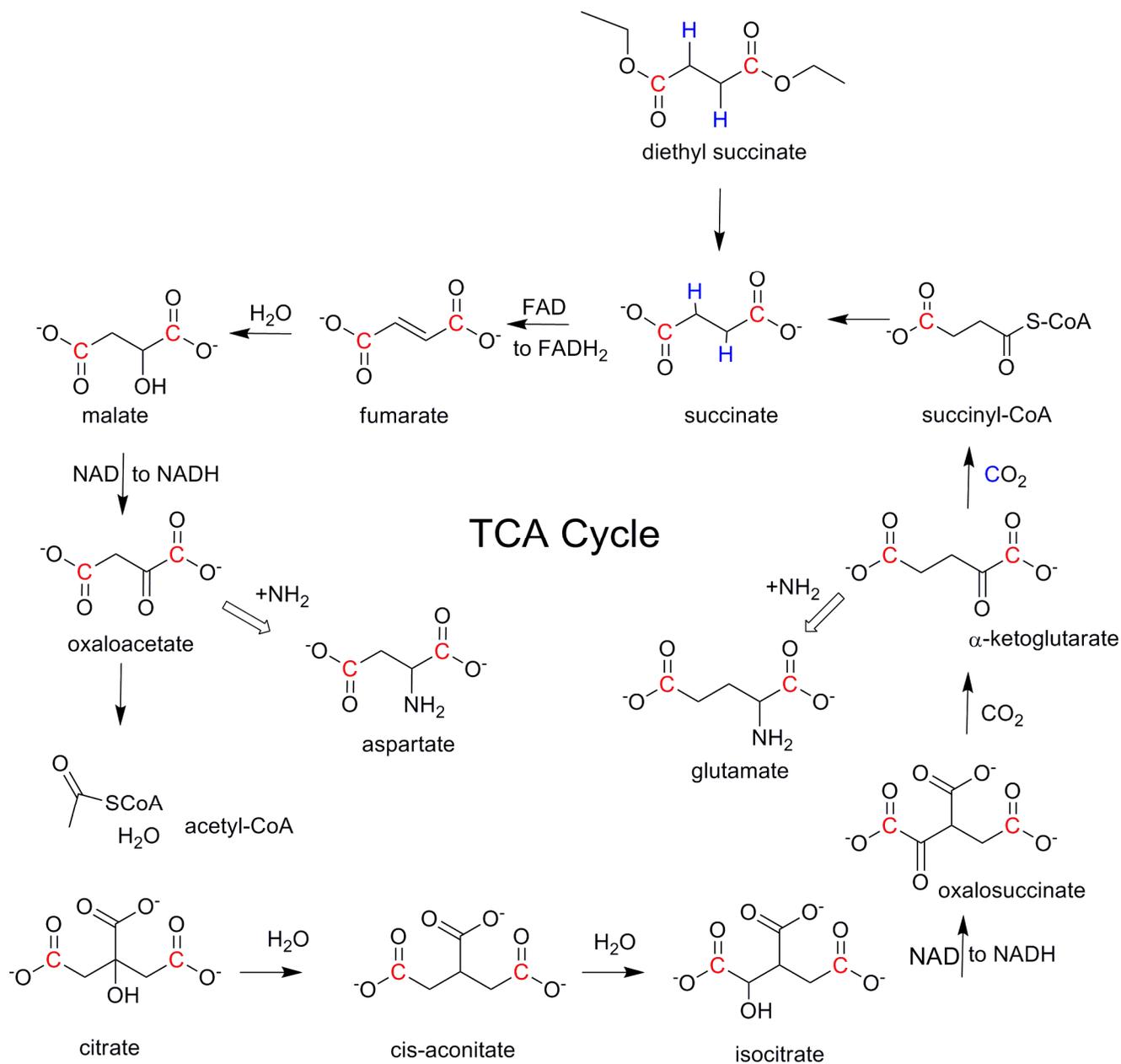
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| Mouse # | Injection type             | Succinate | Malate | Fumarate | Aspartate | Conc. Diethyl succinate |
|---------|----------------------------|-----------|--------|----------|-----------|-------------------------|
| 1       | i.v.                       | X         | X      | X        | X         | 24 mM                   |
| 2       | i.v.                       | X         | X      |          |           | 24 mM                   |
| 3       | i.v.                       | X         | X      |          | X         | 20 mM                   |
| 4       | i.v.                       | X         | X      | X        | X         | 20 mM                   |
| 5       | i.v.                       | X         | X      | X        | X         | 20 mM                   |
| 6       | i.v.                       | X         | X      |          | X         | 20 mM                   |
| 7       | i.v.                       | X         | X      |          | X         | 20 mM                   |
| 8*      | i.p. (1 <sup>st</sup> inj) |           | X      | X        | X         | 20 mM                   |
|         | i.p. (2 <sup>nd</sup> inj) | X         | X      | X        | X         | 20 mM                   |
| 9*      | i.v. (1 <sup>st</sup> inj) | X         | X      | X        | X         | 20 mM                   |
|         | i.p. (2 <sup>nd</sup> inj) |           | X      | X        |           | 20 mM                   |
| 10*     | i.v. (1 <sup>st</sup> inj) | X         | X      | X        | X         | 20 mM                   |
|         | i.p. (2 <sup>nd</sup> inj) | X         | X      | X        | X         | 20 mM                   |
| 12      | i.v.                       | X         | X      | X        | X         | 20 mM                   |
| 13      | i.v.                       | X         | X      | X        | X         | 20 mM                   |
| 14      | i.v.                       | X         | X      | X        | X         | 20 mM                   |

**Supplement Table 1:** Table of Metabolites seen in first injection of hyperpolarized diethyl succinate.

Because we only performed four i.p. injections, the \*asterisks experiments are included in the table.



**Supplement Figure 1:** A diagram representing TCA cycle metabolites arising from hyperpolarized  $1\text{-}^{13}\text{C}$  succinate. Any of the red carbons of the metabolites could be in the hyperpolarized state from the metabolism of hyperpolarized  $^{13}\text{C}$  labeled succinate. Based on our technique, only hyperpolarized metabolites will be seen with *in vivo*  $^{13}\text{C}$  MRS. In our current experiments, we are seeing the left hand side of this diagram (fumarate, malate and aspartate).