

## Supplementary Material

# Noninvasive rapid detection of metabolic adaptation in activated human T lymphocytes by hyperpolarized $^{13}\text{C}$ magnetic resonance

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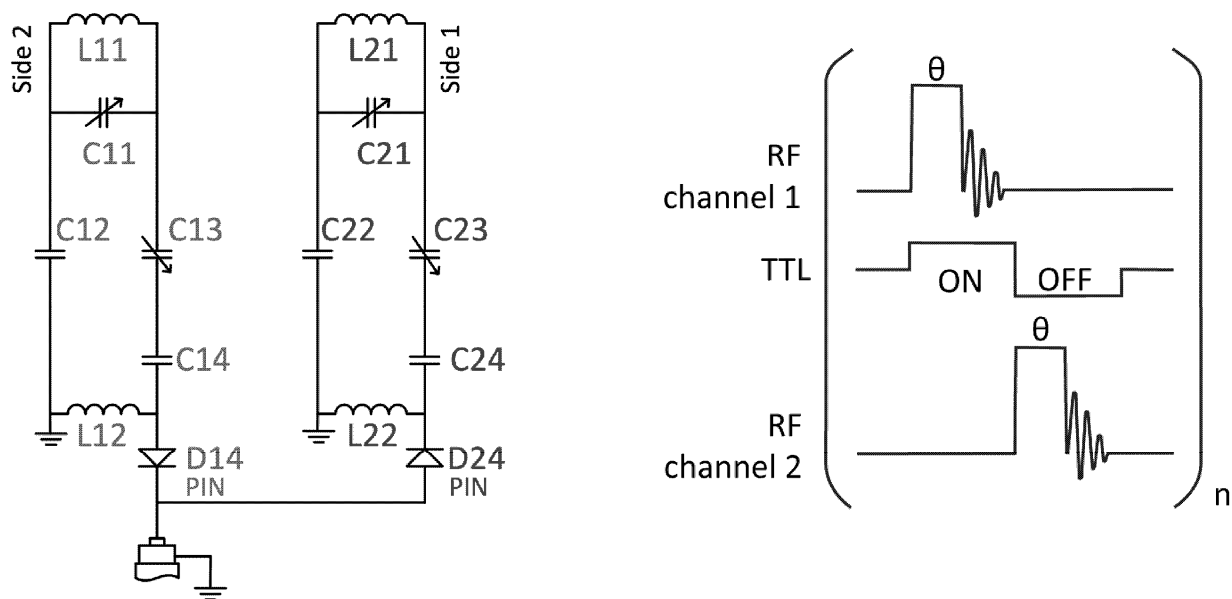
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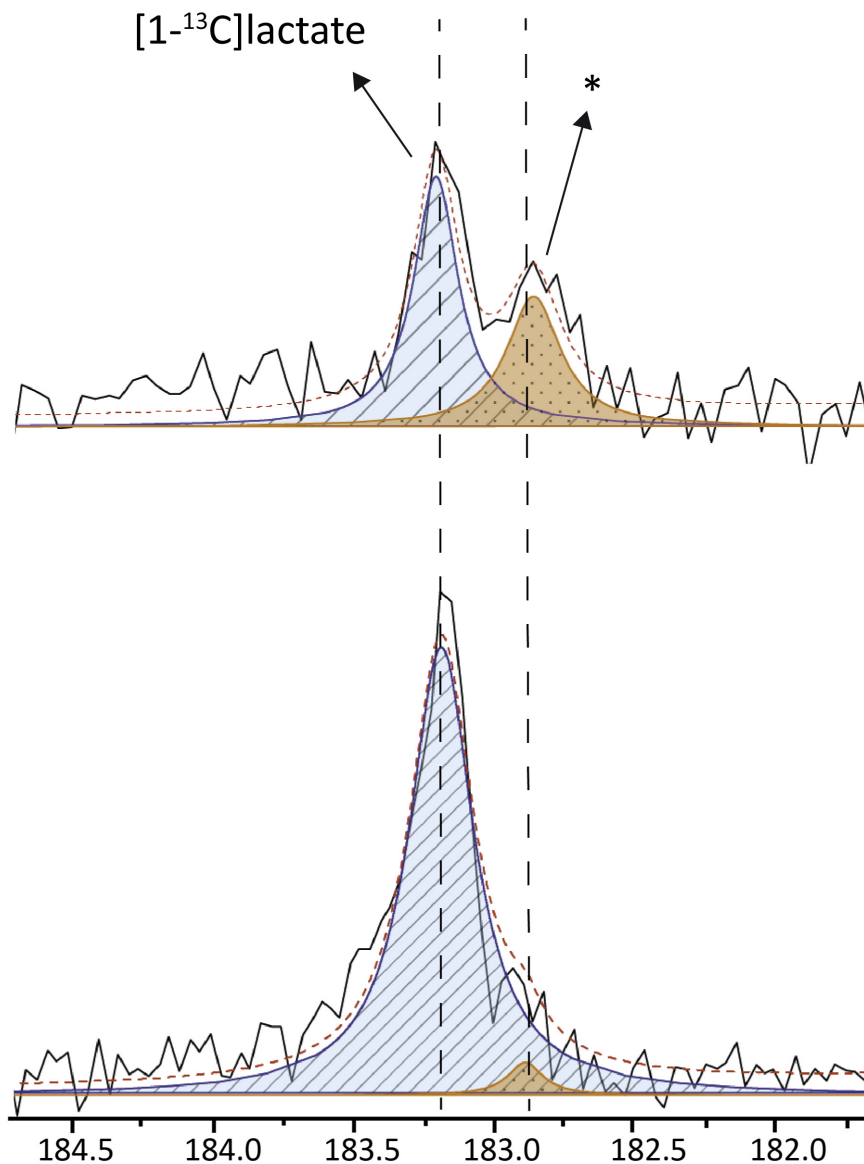
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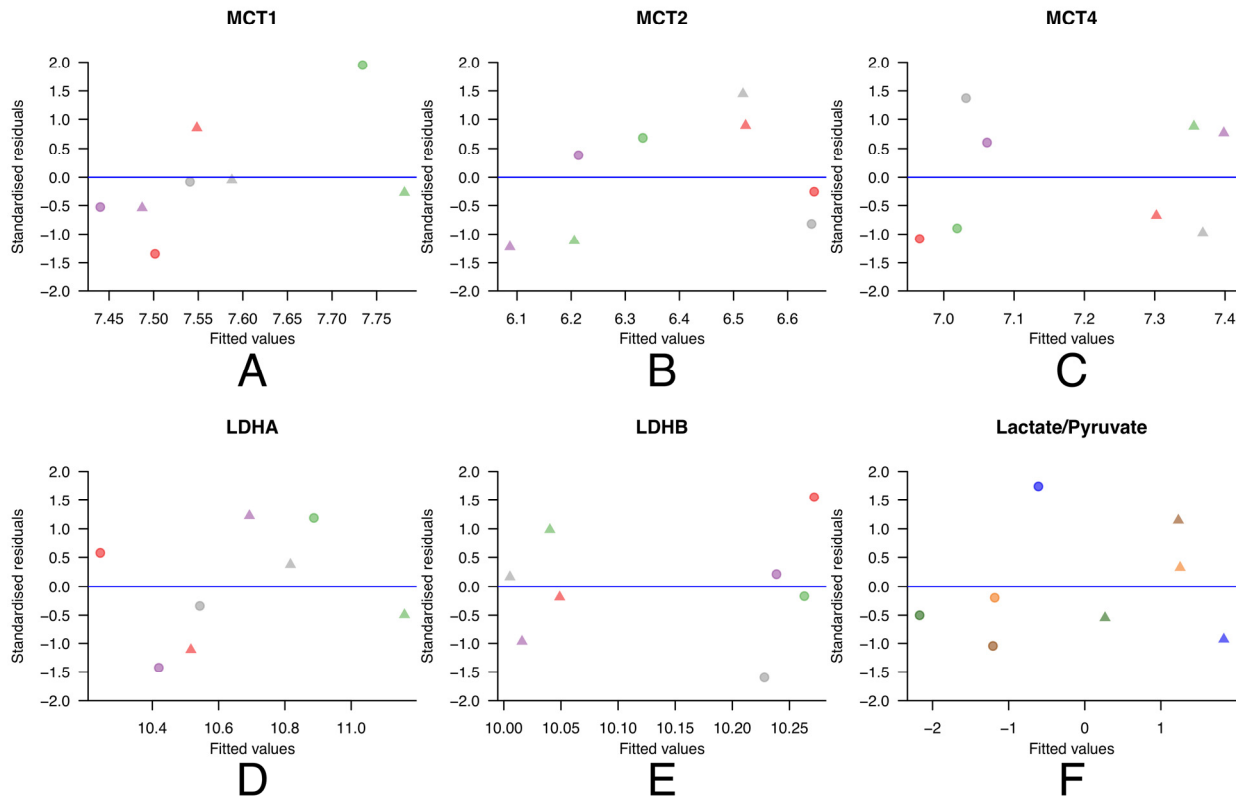
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**Figure S1.** (A) Circuit schematic of one of the two symmetric channels ( $^1\text{H}$  and  $^{13}\text{C}$ ) of the custom-designed MR probe. The circuit scheme is identical for both  $^1\text{H}$  and  $^{13}\text{C}$  channels. The PIN diodes were placed in forward (D24) and reverse (D14) bias to isolate the two probe heads using a transistor-transistor logic (TTL) signal synchronized with the RF pulse via a digitally-controlled driver<sup>1</sup>. The capacitor values used in practice for C12 and C14 are 0.5 pF and 7.5 pF for  $^1\text{H}$  and 6.2 pF and 102.2 pF for  $^{13}\text{C}$ . The capacitor values for the other side of each channel are identical. L11 and L21 represents the inductors of the  $^1\text{H}/^{13}\text{C}$  probe heads. Inductors (L12 and L22, 150nH each) were placed between the PIN diodes and the ground to create a DC current pass through; (B) A schematic diagram showing the pulse sequence implemented for alternating acquisitions of MR signals from each side of the probe (channel 1 corresponds to side 1 and channel 2 to side 2). A total of n acquisitions on each side is preset.



**Figure S2.** Representative spectra showing the impurity signal (marked with a star) overlapping with the lactate peak with a chemical shift difference of 0.3 ppm. To subtract the impurity signal from the [1-<sup>13</sup>C]lactate signal, the peak fitting module from the OriginPro 2019 Peak Analyzer toolbox was used to fit both signals with a fixed chemical shift separation of 0.3 ppm for all experiments. Only the integral of the peak corresponding to the [1-<sup>13</sup>C]lactate signal (light blue hatched area) was used for calculating the lactate-to-pyruvate signal ratios.



**Figure S3.** Residual analysis of the random-intercept linear mixed models fits of mRNA expression of MCTs (A-C), LDHA (D), LDHB (E), and [2,3-<sup>13</sup>C<sub>2</sub>]lactate-to-[2,3-<sup>13</sup>C<sub>2</sub>]pyruvate ratio measured by LC-MS. The plots show the residuals (y-axis) versus the fitted values (x-axis). Point colors correspond to donor and symbols to states: dots for resting and triangles for activated. These model checks, showing symmetry of the residuals around 0 and homoscedasticity, suggest a good fit of the model to the data.

## References

1. Pilloud, Y. & Gruetter, R. in *ESMRMB 2012*. 556-557 (MAGMA).