# Enhancing the Spatial Resolution of Hyperpolarized Carbon-13 MRI of Human Brain Metabolism using Structure Guidance

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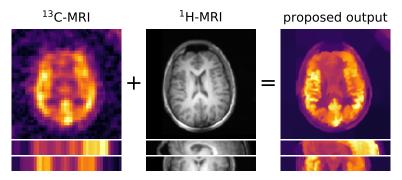
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Engineering and Physical Sciences Research Council

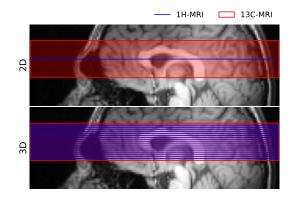


# Motivation



- Dynamic nuclear polarization like Carbon-13 Magnetic Resonance Imaging (<sup>1</sup>3C- MRI) has a relatively low spatial resolution
- enhance 3D resolution of <sup>1</sup>3C- MRI using structural information of Hydrogen-1 MRI (<sup>1</sup>H-MRI)

# Novelty



- Superresolution of 13C-MRI using 1H-MRI gained increased interest recently Farkash et al. MRM 2019, Dwork et al. Magn Res Mat Phys Bio Med 2021, Ma and Park Tomography 2020
- Novelty 1: 2D v 3D anatomical data
- Novelty 2: improved mathematical model (dTV)

#### Inverse problem

$$Sx = y$$

- x: desired high-resolution 13C-MRI image
- y : low-resolution 13C-MRI image
- S : resolution model: 3D high-res to 2D low-res

$$S: \mathbb{R}^{N \times M \times K} \to \mathbb{R}^{N/R \times M/R}, \quad (S \times)_{n,m} = \frac{1}{K \cdot R^2} \sum_{k=1}^{K} \sum_{a,b=0}^{R-1} x_{n+a,m+b,k}$$

Here: K = 15, R = 4

**Goal:** recover 
$$X$$
 given  $Y$ 

## Variational Regularization

**Variational regularization** Approximate solution of Sx = y via

$$\hat{x} = \arg\min_{x} \left\{ \|Sx - y\|_{2}^{2} + \frac{\lambda \mathcal{R}(x)}{\lambda \mathcal{R}(x)} \right\}$$

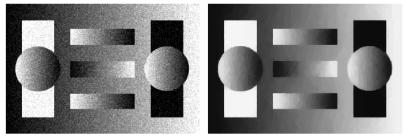
•  $||Sx - y||_2^2$ : data fidelity, can be related to noise statistics

- R: regularizer: penalizes unwanted features, ensures stability and uniqueness
- λ: regularization parameter: λ ≥ 0. If λ = 0, then an original solution is recovered. As λ → ∞, more and more weight is given to the regularizer R.

## Example: Regularizers

► Total Variation Rudin, Osher, Fatemi 1992

$$\mathcal{R}(x) = \|\nabla x\|_{2,1} = \sum_{\beta} \|(\nabla x)_{\beta}\|_2 = \sum_{\beta} \sqrt{(\partial_1 x)_{\beta}^2 + (\partial_2 x)_{\beta}^2 + (\partial_3 x)_{\beta}^2}$$



#### Noisy image

TV denoised image

How to incorporate the 1H-MRI image into the regularization?

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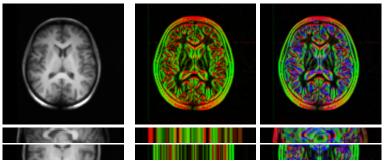
Directional Total Variation Ehrhardt and Betcke 2016

$$\mathcal{R}(x) = \sum_{eta} \|D_{eta}(
abla x)_{eta}\|_2$$

$$D_{eta} = I - \gamma \xi_{eta} \xi_{eta}^T \in \mathbb{R}^{3 imes 3}, \quad \xi_{eta} = rac{(
abla v)_{eta}}{\sqrt{\|(
abla v)_{eta}\|_2^2 + arepsilon^2}} \in [-1, 1]^3 \subset \mathbb{R}^3$$

Default values:  $\gamma \approx 0.9995, \varepsilon \approx 0.01 \cdot \max_{\beta} \| (\nabla \nu)_{\beta} \|_2$ 

#### 2D v 3D directional total variation <sup>1</sup>H-MRI vector field $\xi$ (2D) vector field $\xi$ (3D)



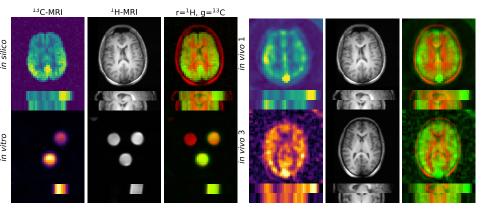
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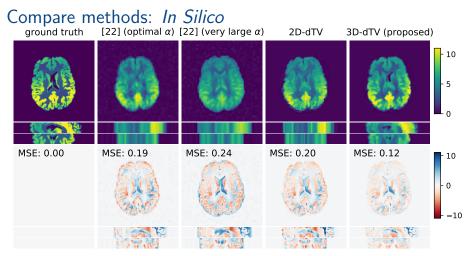
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## Overview of Data

- ► In vitro 3 tubes Daniels et al. NMR Biomed 2016
- ► In vivo 4 healthy volunteers Grist et al. NeuroImage 2019
- In silico GM/WM  $\approx$  4, smooth variations



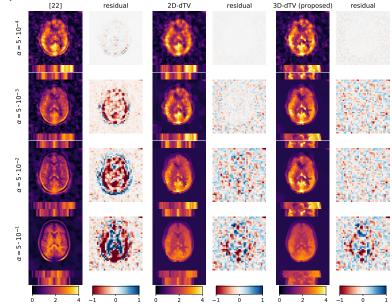
Images need to be registered!



► [22] = Dwork et al. 2021 enforces <sup>13</sup>C-MRI and <sup>1</sup>H-MRI globally to either have positive or negative correlation (see CSF for large  $\alpha$ )

 3D-dTV leads to anatomically better-defined structures compared to 2D-dTV

## Compare methods: In Vivo

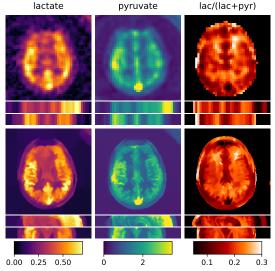


Similar observations as for in silico data

# lactate pyruvate lac/(lac+pyr) 0.00 0.25 0.50 0.0 0.5 1 2 0

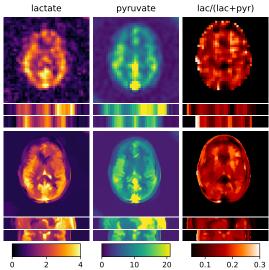
Higher resolution but preserves smooth variations

# Qualitative evaluation: In Vitro



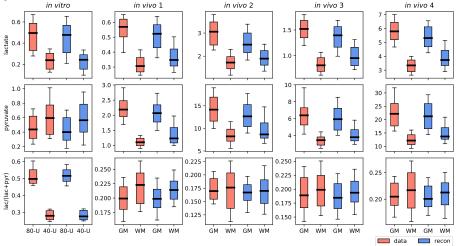
Anatomically well-defined images

not constant within anatomical regions (e.g. GM, WM)



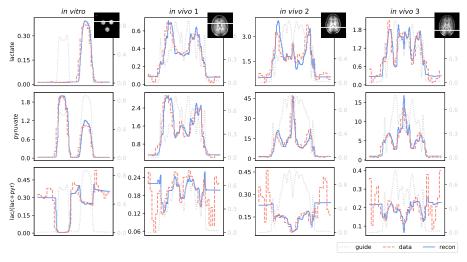
**Similar observations** as for the 1st *in vivo* data set

## Quantitative evaluation



- Similar mean values in low and high-resolution <sup>13</sup>C-MRI images
- Variation within and difference between anatomical regions reduced (good? bad? just reflecting truth?)

# Line plots



Further visual evidence that quantification is preserved

# Conclusions and Outlook

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- ► Aim: increase resolution of <sup>13</sup>C-MRI
- Directional total variation well-suited for this task (allows locally changing correlations)
- 3D guide image works much better than 2D as it better reflects anatomy
- Higher resolution with preserved quantification
- Computationally efficient: takes less than 2 minutes on 3 year old MacBook Pro
- Robust to parameter choices: simple to tune the 3 model parameters and 2 algorithm parameters

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#### Future work

- Validation, validation, validation ... e.g. in vivo data sets with tissue samples
- Better modelling of data, e.g. by using k-space data

