

Metabolic Imaging of Traumatic Brain Injuries Using Ultrahigh-Resolution ^1H -MRSI

Tianyao Wang¹, Jun Liu¹, Tianxiao Zhang², Ziyu Meng², Danni Wang², Ke Xue², Yudu Li^{3,4}, Rong Guo^{3,4}, Yibo Zhao^{3,4}, Xin Yu⁵, Zhi-Pei Liang^{3,4}, and Yao Li²

¹Radiology Department, The Fifth People's Hospital of Shanghai, Fudan University, Shanghai, China, ²Institute for Medical Imaging Technology, School of Biomedical Engineering, Shanghai Jiao Tong University, Shanghai, China, ³Beckman Institute for Advanced Science and Technology, University of Illinois at Urbana-Champaign, Urbana, IL, United States, ⁴Department of Electrical and Computer Engineering, University of Illinois at Urbana-Champaign, Urbana, IL, United States, ⁵Department of Biomedical Engineering, Case Western Reserve University, Cleveland, OH, United States

Synopsis

Traumatic brain injury (TBI) is a significant public health problem that contributes to a large number of injury-induced deaths each year. MRSI has long been recognized as a powerful tool for detection of neurometabolic alterations induced by TBI; however, most existing MRSI studies of TBI are limited by low resolution which severely reduce the detection sensitivity. In this study, we performed MRSI scans on TBI patients using a newly developed ultrahigh-resolution ^1H -MRSI technique. Our experimental study yielded very encouraging results and showed that ultrahigh-resolution MRSI can capture neurometabolic alterations induced by TBI effectively.

Introduction

MRI studies of traumatic brain injury (TBI) have grown significantly over the past decade. Many advanced MRI methods, such as diffusion tensor imaging (DTI), susceptibility-weighted imaging (SWI), perfusion-weighted imaging, and fMRI, are being used to detect many of the functional and pathophysiological alterations resulting from TBI, and have provided significant insights and understanding of TBI^{1,2}. MR spectroscopic imaging (MRSI) has long been recognized as an ideal tool for detection of neurometabolic alterations induced by TBI³. However, clinical and research applications of whole brain ^1H -MRSI of TBI have been extremely limited due to several long-standing technical barriers, including long data acquisition time, poor spatial resolution, low SNR and overwhelming background water and lipid signals. As a result, most existing MRSI studies of TBI were carried out in low resolution (e.g., $10\times 10\times 7.5\text{ mm}^3$ nominal resolution) or using single-voxel techniques³. The partial volume effects associated with low-resolution/single-voxel techniques can “dilute” abnormal metabolic changes and reduce the sensitivity for detection; it has also been recognized that these techniques cannot distinguish focal from diffuse injury. In this study, we performed ^1H -MRSI experiments on TBI patients using a newly developed ultrahigh-resolution MRSI technique known as SPICE. The latest version of SPICE can cover the whole brain of $240\times 240\times 120\text{ mm}^3$ with $2.0\times 2.4\times 2.0\text{ mm}^3$ nominal resolution in a 10-minute scan. Our experimental study yielded very encouraging results and showed that ultrahigh-resolution MRSI can capture neurometabolic alterations induced by TBI effectively.

Method

The in vivo scans were performed on a 3T MR scanner (Siemens Skyra) with IRB approved at the Fifth People's Hospital of Shanghai, China. The MRSI data were acquired using the latest version of SPICE (SPectroscopic Imaging by exploiting spatiospectral CorrElation)^{4,5}. The data acquisition scheme has the following novel features: a) FID acquisitions with an ultrashort-TE (1.6 ms) and very-short-TR (160 ms), b) no water/lipid suppression, c) variable density sampling of (k, t)-space, d) rapid and extended k-space coverage with EPSI trajectories, and e) incorporation of navigators for detection and correction of field drift and subject head motion. The data acquisition scheme achieves a nominal resolution $2.0\times 2.4\times 2.0\text{ mm}^3$ with whole brain coverage (FOV: $240\times 240\times 120\text{ mm}^3$) in a 10-minute scan.

Reconstruction of the spatiospectral function from the SPICE data is accomplished using a union-of-subspaces model, incorporating pre-learned spectral basis functions^{4,5,7}. These spectral basis functions include the resonance structures of the detectable molecules (generated using quantum mechanical simulations) and their lineshape functions pre-learned from training data. Spectral quantification is done using an improved LCmodel-based algorithm that incorporates both spatial and spectral priors⁹. QSM is extracted from the water spatiospectral information⁶.

To identify the injured region, diffusion-weighted imaging (DWI) (TR/TE = 11200/106 ms, resolution = $2\times 2\times 2\text{ mm}^3$, FOV = 256 mm, 70 slices, b = 1000/2000/3000 s/mm², 30 directions for each b value and 12 b = 0 images), 3D MPRAGE imaging (TR/TE/TI = 2500/2.26/900 ms, resolution = $1.0\times 1.0\times 1.0\text{ mm}^3$, FOV = 256 mm, 176 slices) and Fluid-Attenuated Inversion Recovery (FLAIR) imaging (TR/TE = 9000/89 ms, resolution = $0.5\times 0.5\times 2.0\text{ mm}^3$, FOV = 240 mm, 82 slices) were performed in the scan session. The regions of interest were identified from the Trace and fiber orientation distribution (FOD) images calculated from the DWI image using Syngo workstation and MRTRIX3 software⁸.

Results and Discussion

Figure 1 shows the high-resolution metabolic maps (includes N-acetylaspartate (NAA), creatine (Cr), choline (Cho) and myo-inositol (ml)) reconstructed from the ^1H -MRSI data, which cover the whole brain (FOV: $240\times 240\times 120\text{ mm}^3$). Figure 2 shows Trace image and FOD image as well as the QSM and NAA maps obtained from the MRSI data. As can be seen, the large cerebral contusion lesion can be identified from the QSM map. The NAA reduction is shown in regions of both the large cerebral contusion lesion and the small diffuse axonal injury (DAI) lesion. The spatially resolved spectra are presented in Fig. 3. A reduction in NAA can be observed from the spectra in both cerebral contusion lesion and DAI lesion and the reduction is more significant in the cerebral contusion region. The capability to detect the millimeter DAI lesion is of great importance in TBI metabolic study.

Conclusions

We have successfully performed in vivo metabolic imaging experiments on TBI patients using ultrahigh-resolution ^1H -MRSI. Our experimental results show that neurometabolic changes in both cerebral contusion lesions and small size DAI lesions can be captured by the high-resolution metabolite maps and its corresponding localized spectra. Our study may lay the foundation for further investigation and application of ultrahigh-resolution metabolic imaging for studying neurometabolic alterations in TBI.

Acknowledgements

This work is supported by National Science Foundation of China (No.61671292 and 81871083) and shanghai municipal commission of health and family planning (No. 20154Y0094)

References

1. Wu X, Kirov I, Gonen O, et al. MR imaging applications in mild traumatic brain injury: an imaging update. *Radiology* 2016, 279: 693-707.
2. Cook GA, Hawley JS. A review of mild traumatic brain injury diagnostics: current perspectives, limitations, and emerging technology. *Military medicine* 2014, 179:1083-1089.
3. Lin AP, Liao HJ, Merugumala S K, et al. Metabolic imaging of mild traumatic brain injury. *Brain Imaging Behav* 2012, 6:208-223.
4. Lam F and Liang ZP. A subspace approach to high-resolution spectroscopic imaging. *Magn Reson Med* 2014, 71:1349-1357.
5. Lam F, Ma C, Clifford B, et al. High-resolution ^1H -MRSI of the brain using SPICE: data acquisition and image reconstruction. *Magn Reson Med* 2016, 76:1059-1070.
6. Peng X, Lam F, Li Y, et al. Simultaneous QSM and metabolic imaging of the brain using SPICE. *Magn Reson Med* 2018, 79:13-21.
7. Ma C, Lam F, Johnson CL, et al. Removal of nuisance signals from limited and sparse ^1H MRSI data using a union-of-subspaces model. *Magn Reson Med* 2016, 75:488-497.
8. Tournier JD, Calamante F, Connelly A. MRtrix: diffusion tractography in crossing fiber regions. *Int J Imag Syst Tech* 2012, 22:53-66.
9. Li Y, Lam F, Clifford B, et al, A Subspace Approach to Spectral Quantification for MR Spectroscopic Imaging. *IEEE Trans Biomed Eng* 2017, 64:2486-2489.

Figures

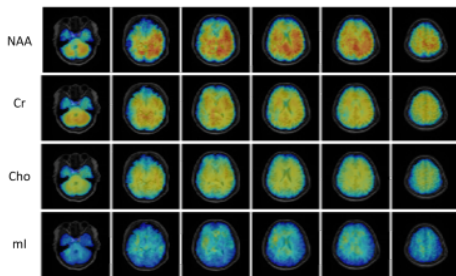


Figure 1. Reconstructed high-resolution metabolite maps including NAA, Cr, Cho and ml. The FOV covers the whole brain ($240 \times 220 \times 120 \text{ mm}^3$) and the MRSI acquisition takes 10 minutes.

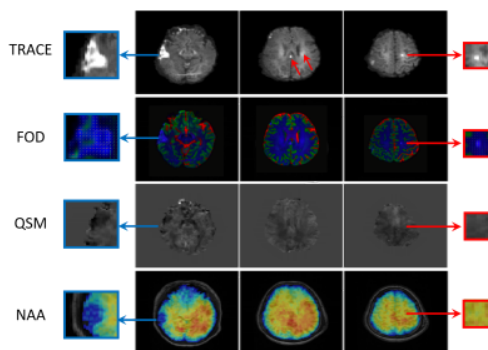


Figure 2. The Trace DWI, FOD, QSM and NAA maps of the slices where both the cerebral contusion lesion and diffuse axonal injury (DAI) lesion exist. Both the cerebral contusion and small DAI lesions can be identified from the NAA map where the NAA concentration is lower than in the surrounding tissues. The large cerebral contusion lesion can also be identified from the QSM image.

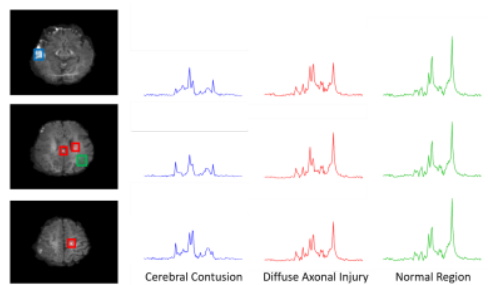


Figure 3. Representative spectra in the cerebral contusion region, diffuse axonal injury region and the normal tissue region. The reduction in NAA can be observed from both the cerebral contusion region and the diffuse axonal injury region.