

PET-MR compatible CEST method for imaging of Alzheimer's Disease

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INTRODUCTION:

PET scans are the gold standard for detecting amyloid deposits in the brain¹. CEST imaging may provide alternative or complementary information to PET imaging. However, typical CEST experiments utilizes long RF saturation that is not compatible with the PET-MR hybrid system. The aim of this study is to address this limitation by developing a steady-state CEST MRI method that allows CEST imaging to be acquired during PET sessions.

METHODS:

Our CEST sequence incorporates steady-state RF saturation with stack-of-stars golden-angle radial sampling². There is no delay between the acquisitions, ensuring the attainment of a steady-state saturation condition. Gaussian pulses were used for CEST labeling with a B1 = 0.5 μ T, duration of 50 ms, followed by a spoiling period (20 mT/m, 5 ms). Images were acquired at 32 saturation frequency offsets between -8.1 to 3.0 ppm, along with two reference images. 100 radial stacks were acquired at each frequency offset, corresponding to a total scan time of 5.5 min. The study was approved by the Mount Sinai institutional review board and was conducted on individuals with mild cognitive impairment and age matched controls. GRASP-Pro technique was performed to reconstruct the dynamic images³. rNOE Lorentzian difference (LD_{noe}) maps were generated by computing Lorentzian difference within the range of -3.2 to -3.8 ppm. The Δ ST(-3.6 ppm) maps⁴ were calculated by taking the difference between the signal at -8 and -3.6 ppm. Amyloid PET was acquired using 18F-labeled florbetaben (Neuraceq). The amyloid SUVR map was calculated using cerebellum gray matter as reference region. Both CEST and PET images were co-registered to the anatomical T₁ images.

RESULTS:

Two ROIs were delineated, representing the white matter (WM) and the gray matter (GM) areas. Fig. 1A shows the full z-spectra in the GM and WM. A Lorentzian line was fitted to the negative side the z-spectrum (dotted line in Fig 1B) and rNOE effect can be isolated by taking the difference between the Lorentzian fitted spectrum and the acquired z-spectrum (Fig. 1C). Fig. 2. shows T1-weighted, LD_{noe}, Δ ST(-3.6 ppm) and amyloid SUVR maps of an A β negative, and an A β positive individual, respectively.

DISCUSSION:

The method can provide excellent rNOE contrast, potentially sensitive to beta-amyloid aggregates. Compared to healthy controls, the AD/MCI patients appeared to have narrower distributions of LD_{noe} and Δ ST and a trend for higher values. Ongoing work includes increase subject accrual and correlation of the CEST and the A β PET.

CONCLUSION:

We demonstrated the implementation of a steady-state, radially acquired imaging sequence that is fully compatible to the hybrid PET-MR system. The method is potentially sensitive to A β deposition, but more data is needed to validate the findings.

ACKNOWLEDGMENTS:

Ms. Sarah Binder and Ms. Nancy Andrew-Jaja for assisting with participant recruitments. Drs Johanne Bezy-Wendling and Fanny Noury (University of Rennes, LTSI-Inserm U1099, 35000 Rennes, France) for their support. Funding support from the NIH: R00EB026312, R21AG083685

REFERENCES: (1) Vallabhajosula, S. et al *Seminars in nuclear medicine*, 2011; Elsevier: Vol. 41, pp 283-299. (2) Ma, S.; et al. *Magnetic resonance in medicine* **2022**, 87 (1), 102-119. (3) Feng, L.; et al. *Magnetic resonance in medicine* **2020**, 83 (1), 94-108. (4) Chen, L.; et al. *Neuroimage* **2019**, 188, 380-390.

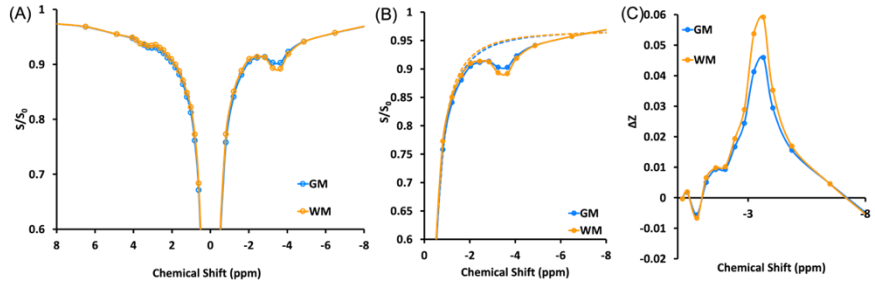


Fig. 2. (A) The z-spectrum acquired with the ST MRI sequence in the gray and white matter regions; (B) The negative side of the acquired z-spectrum and the Lorentzian fitted spectrum; (C) the Lorentzian difference spectrum.

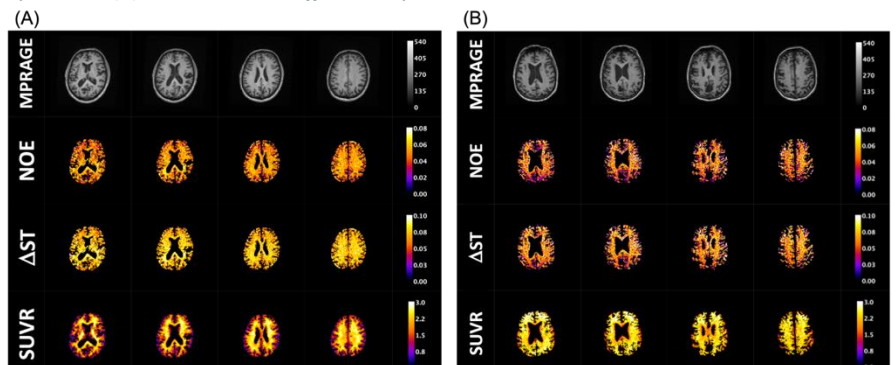


Fig. 1. From top to bottom: Co-registered T1 MPRAGE, LD_{noe} maps, Δ ST(-3.6 ppm) maps and the A β SUVR maps for (A) an A β negative individual and (B) an A β positive individual.