Program No. 4375



## T2 Relaxometry Based CSF Fraction (CSFF) Mapping is a Better Biomarker for Brain Drainage Pathology Than DTI-based Free Water (DTI-FW) Mapping

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## Declaration of Financial Interests or Relationships

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I have no financial interests or relationships to disclose with regard to the subject matter of this presentation.

# Hypothesis



- The perivascular space (PVS) is filled with cerebrospinal fluid (CSF)-like free water.
- PVS plays a role of pathway for the clearance of metabolites. Enlarged PVS due to blockage of CSF will damage the clearance function.
- CSF fraction map measured by T2-relaxometry is a biomarker of PVS. DTI-FW contains more than CSF free water.

## Program No. 4375 CSF fraction VS DTI-FW Methods: T2 relaxometry based water mapping (CSFF)



• The total water in the brain was modeled as a three-compartment model:

$$S(TE) = A_{my}e^{-TE/T_{2,my}} + A_{ie}e^{-TE/T_{2,ie}} + A_{csf}e^{-TE/T_{2,csf}}.$$
 [1]

where  $A_{my}$ ,  $A_{ie}$ ,  $A_{csf}$  are components for myelin, intro-extra, and CSF water.

• Data fitting can be done using nonlinear least square:

$$\boldsymbol{x} = argmin_{\boldsymbol{x}} \sum_{n=1}^{N} \left| \left| S(\boldsymbol{x}, TE_n) - S_{measure}^n \right| \right|_2^2 + \lambda \left| \left| \nabla_s \boldsymbol{x} \right| \right|_2^2$$
 [2]

• The free water (cerebrospinal fluid fraction, CSFF) is defined as:

$$CSFF = A_{csf} / (A_{my} + A_{ie} + A_{csf}) * 100.$$
 [3]

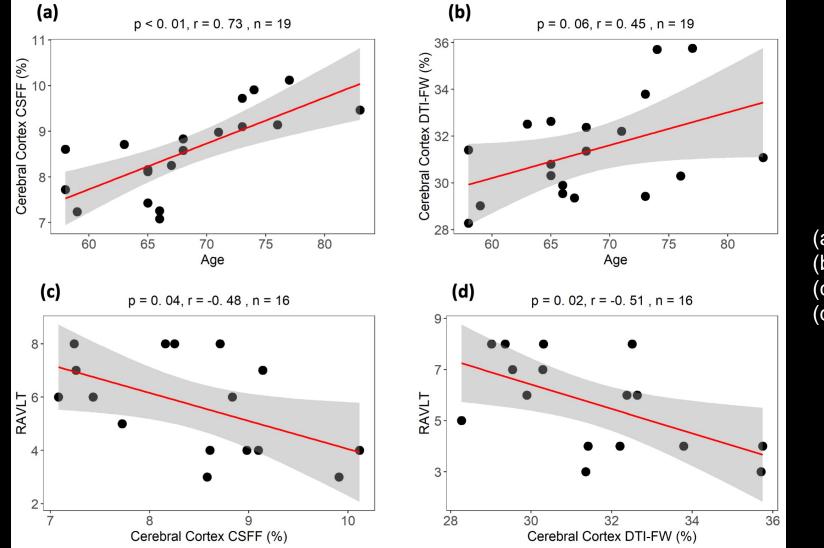
# Methods: Subjects



- 19 subjects age ranges from 55 to 82 years old were scanned with MRI FAST-T2, DTI, PC-MRI, among those 16 have done cognitive test, 8 received PiB PET scan and 10 received MK6240 PET scan.
- Multi-echo T2 data was acquired with Fast Acquisition with Spiral Trajectory and adiabatic T2prep (FAST-T2) sequence at 3T. TEs = 0, 7.6, 17.6, 67.6, 147.6, 307.6 ms. Corresponding T1w, T2w, and T2FLAIR were also acquired at the same session for the anatomical structure and disease diagnosis.

## Results: Free water vs age, and cognitive function

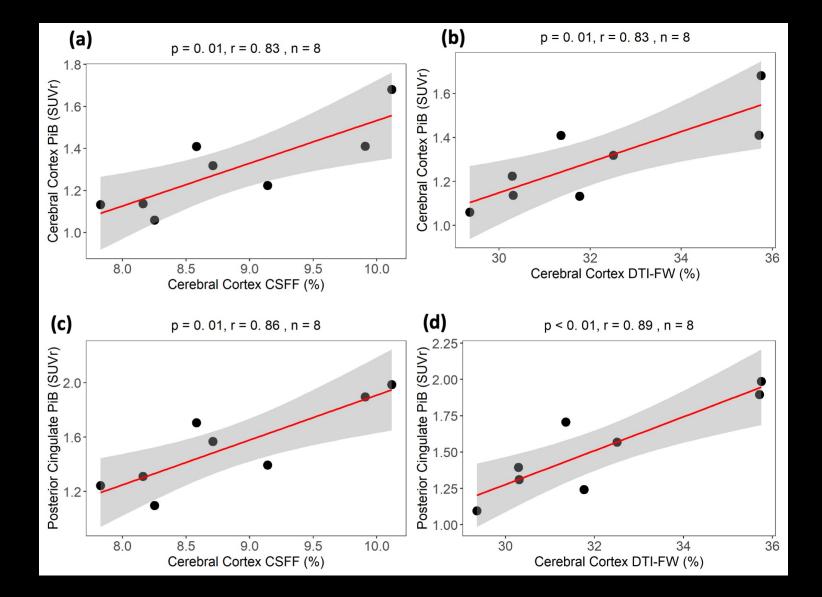




- (a) CSFF vs Age (b) DTI-FW vs Age
- (c) CSFF vs Cognitive function
- (d) DTI-FW vs Cognitive function

### **Results: Free water with A-beta deposit**

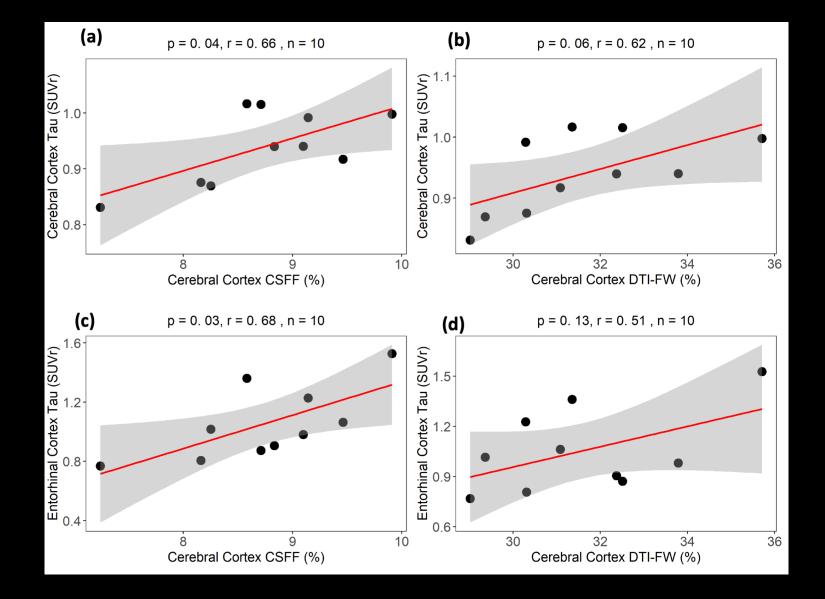




(a) CSFF vs GM A-beta deposit
(b) DTI-FW vs GM A-beta deposit
(c) CSFF vs PC A-beta deposit
(d) DTI-FW vs PC A-beta deposit

## Results: Free water with tau deposit

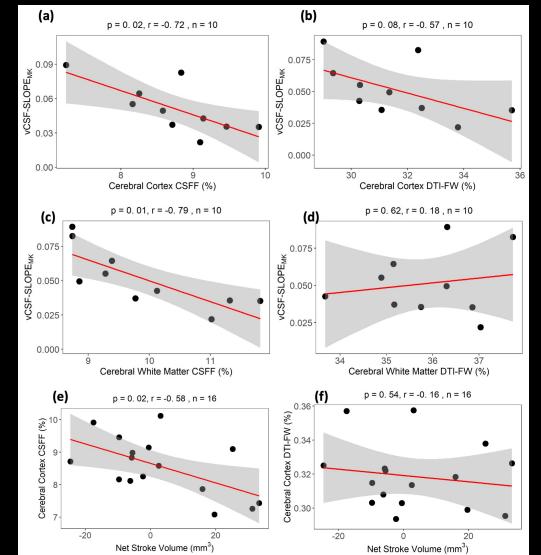




(a) CSFF vs GM tau deposit
(b) DTI-FW vs GM tau deposit
(c) CSFF vs Ent tau deposit
(d) DTI-FW vs Ent tau deposit

#### Program No. 4375 CSF fraction VS DTI-FW Results: Free water with brain clearance slope, and aqueduct CSF flow





- (a) GM CSFF vs vCSF slope
- (b) GM DTI-FW vs vCSF slope
- (c) WM CSFF vs vCSF slope
- (d) WM CSFF vs vCSF slope
- (e) CSFF vs stroke volume
- (f) DTI-FW vs stroke volume



#### **Results: summary**

		Age	RAVLT Score	vCSF- Slope MK6240	Net Stroke Volume	Cerebral Cortex PiB (SUVr)	Posterior Cingulate (SUVr)	Cerebral Cortex Tau (SUVr)	Entorhinal Cortex Tau (SUVr)
Cerebral cortex CSFF	р	<0.01	.0.04	0.02	0.02	0.01	<0.01	0.04	0.03
	r	0.73	-0.48	-0.72	-0.58	0.83	0.86	0.66	0.68
	R <sup>2</sup>	0.50	0.22	0.47	0.28	0.64	0.68	0.36	0.39
Cerebral Cortex DTI- FW	р		0.02			0.01	<0.01		
	r		-0.51			0.83	0.89		
	R <sup>2</sup>		0.29			0.64	0.74		
Cerebral white matter CSFF	р			0.01					
	r			-0.79					
	R <sup>2</sup>			0.57					
Cerebral white matter DTI- FW	р								
	r								
	R <sup>2</sup>								

## **Conclusions**



- Multi-echo spiral T2 relaxometry provides a way to quantify the parenchymal CSF free water.
- CSFF is more sensitive than DTI-FW in all the tested measures, including age, cognitive function, A-beta deposit, tau deposit, vCSF and stroke volume.
- CSFF can be a new biomarker to monitor the glymphatic clearance function and may help us better understand the neurodegenerative diseases.



# Thank you for watching.

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