

Towards mapping CSF volume change at high resolutions at 7T

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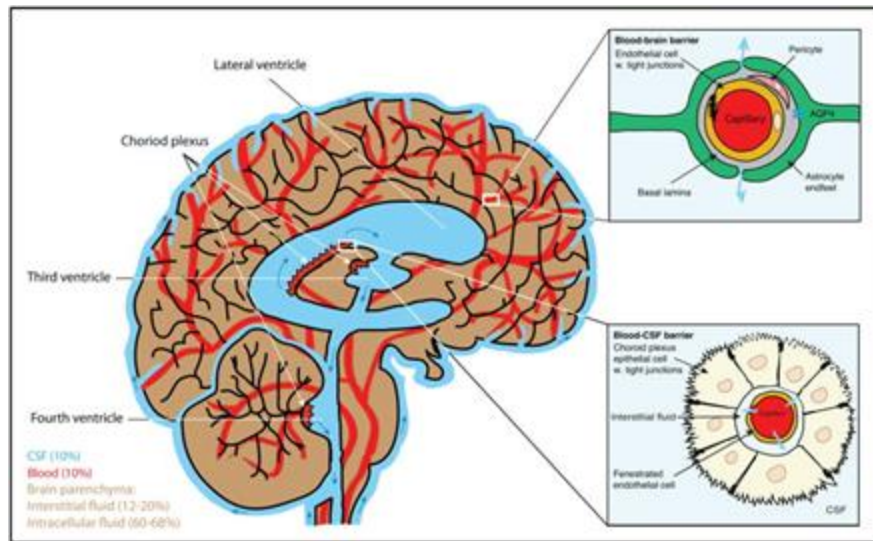


Declaration of Financial Interests or Relationships

Speaker Name: Stephanie Swegle

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

Background on studying cerebrospinal fluid (CSF)



fluid compartments of the brain (Jessen, 2015)

- CSF removes waste and toxins in the brain, especially during sleep
- CSF *flow*: commonly used (Fultz, 2019; Kim, 2022; Grubb, 2019; Gonzalez-Castillo, 2022)
 - but challenged by low velocities, limited imaging coverage, and directional dependencies
 - *Volume* may be more straightforwardly measured
- Contributor and contaminant in VASO (Scouten & Constable, 2008)

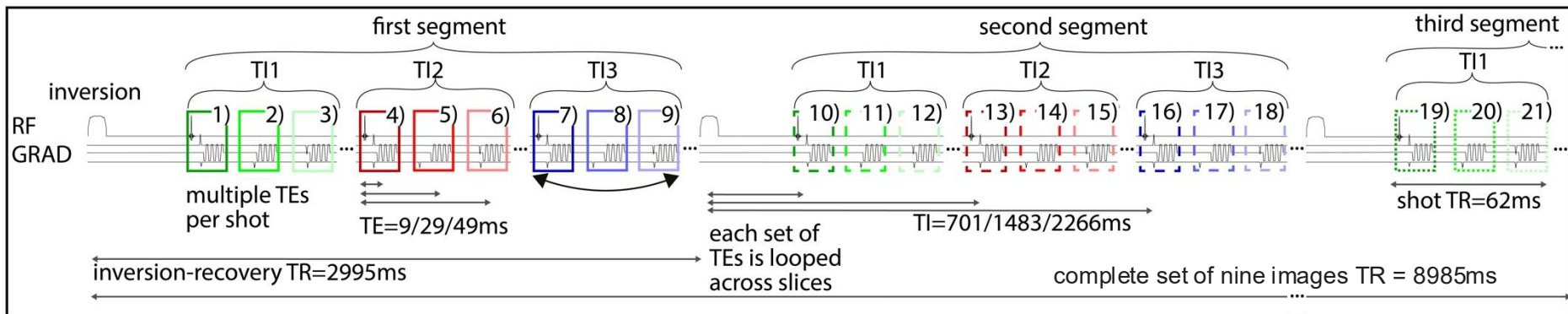
We use a VASO-like inversion recovery (IR) sequence to capture CSF volume changes, independent of dynamic changes in CBV and BOLD

Methods

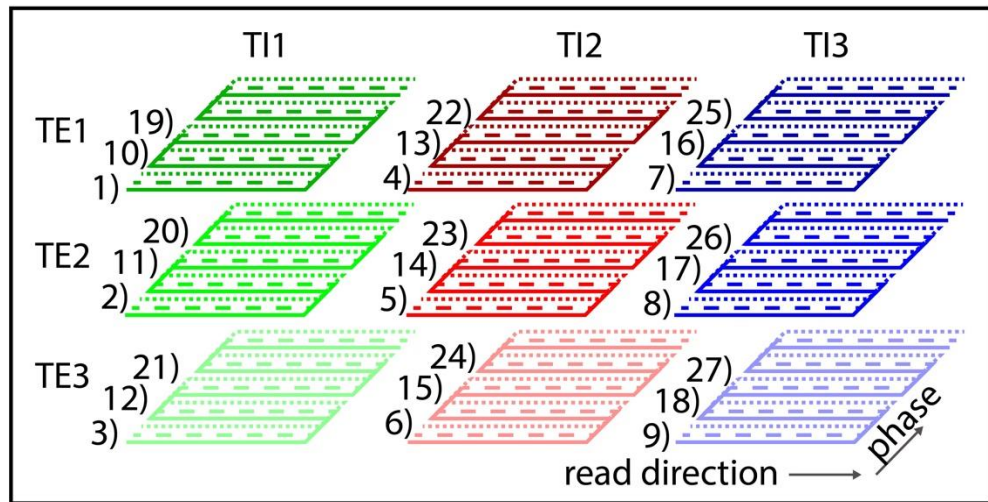
- 7T (N=12 sessions at 7T (Siemens, Germany), 8Tx/32Rx head coil (Nova, USA))
- Multi-echo, multi-inversion
- 3D-EPI with Skipped-CAIPI (Stirnberg, 2021)
- Representative values:
 - TE1/TE2/TE3=9/29/49ms
 - TI1/TI2/TI3=731/1573/2416ms
- Bloch modeling (Huber, 2014; Ivanov, 2013; Donahue, 2011):
 - Assumed T1 values GM/blood/CSF = 1950/2100/4000ms
 - Assumed T2* values GM/blood/CSF/WM = 33.2/37.5/4000/26.815ms
- 12-14 minute block design (30sec blocks) with flashing checkerboards
- Physiological recording to track changes in respiration (Aktas, 2019) (BIOPAC) and in-scanner eye tracking (EyeLink-1000Plus, SR-Research) were done simultaneously to fMRI
- Sample scan parameters (high resolution): 0.8mm iso, 12 slices, $TR_{IR}=3s$, $TR_{vol}=8.8s$
- Sample scan parameters (whole brain **GRAPPA 21**): 2.6mm iso, 80 slices, $TR_{IR}=2.5s$, $TR_{vol}=2.5s$



3D-EPI sampling scheme, segmentation across IR-cycles and 2 phase encoding dimensions



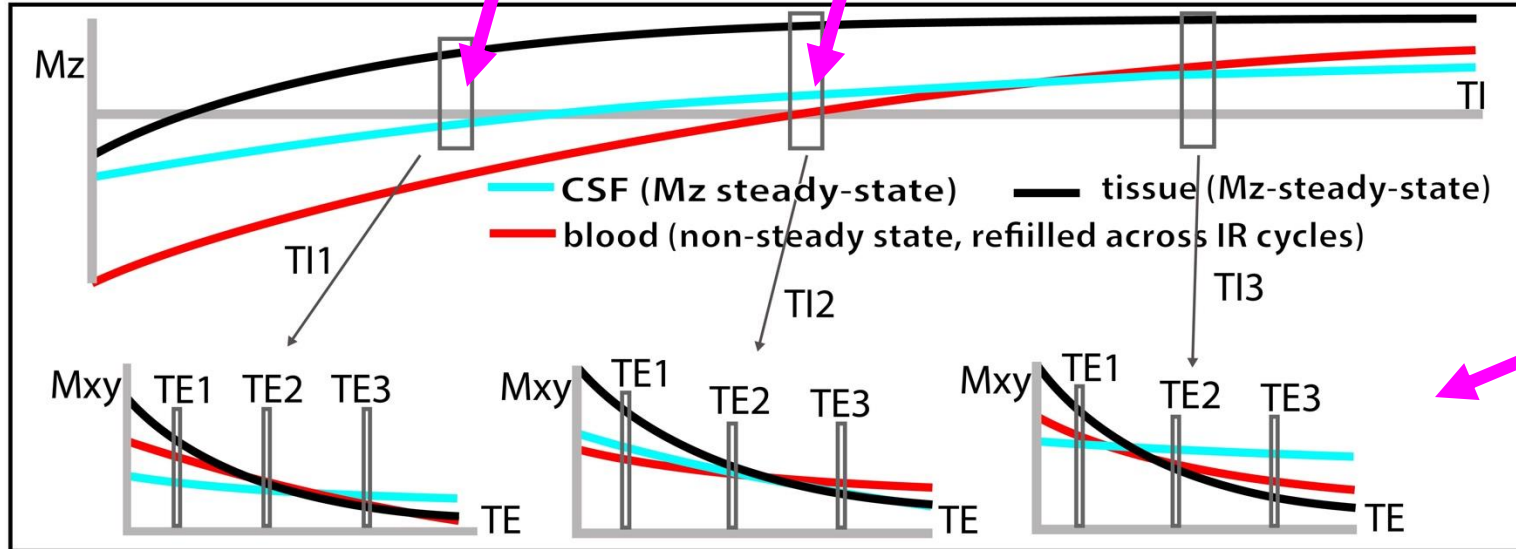
Grouping of 27 shots,
sub-sampled data to
nine echo planar k-
space sets/volumes



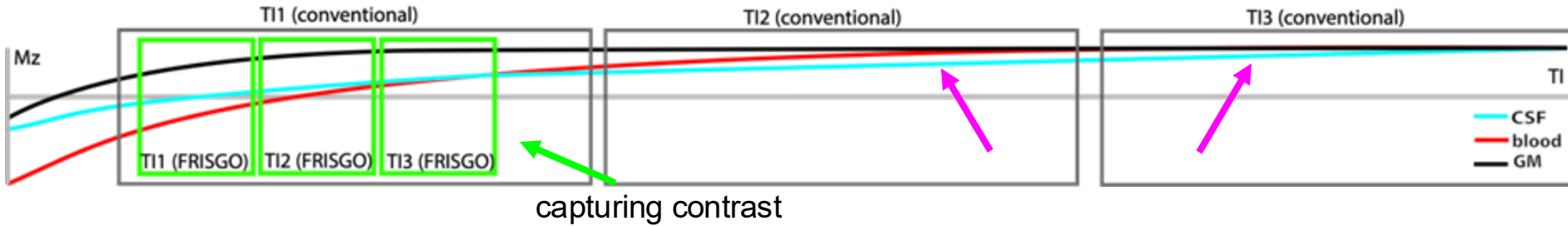
Using CSF's specific relaxation “fingerprint” to separate CSF from CBV and BOLD

CSF is nulled at around $TI1$

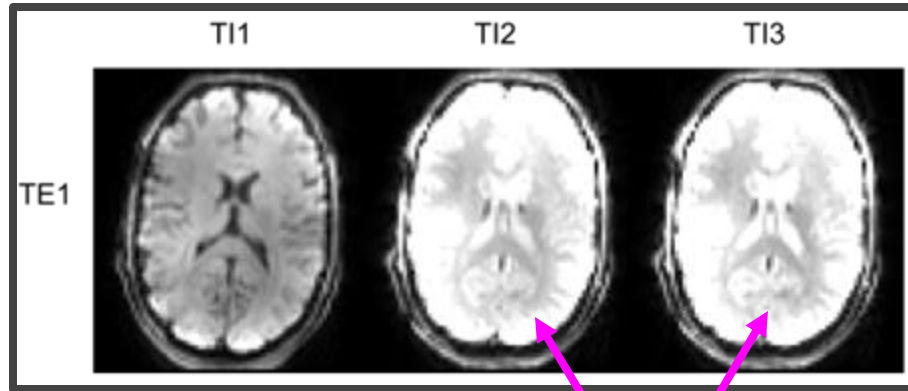
CBV is nulled at around $TI2$



Segmentation across IR cycles + advanced readout acceleration (FRISGO, Huber et al., **#1262**) is needed to capture dynamics of inversion recovery

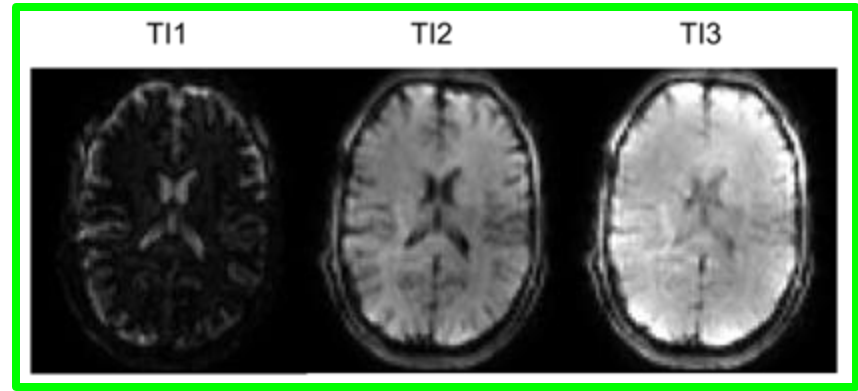


Conventional EPI is not fast enough to fully capture multi-exponential T1 relaxation



no meaningful T1 contrast

FRISGO (Huber et al., **#1262**) can capture multiple whole brain multi-echo images in time scales of T1



Now, can capture the relevant T1-relaxation in the range of T_i s that separates CSF and blood

Bloch Modeling

$$\begin{aligned}
S(Tl_i, TE_i, vol) = & \\
CSF_{vol} * & \left[1 - e^{-Tl_i/T_{1,CSF}} - (M_{CSF}(t=0))e^{-Tl_i/T_{1,CSF}} \right] e^{-TE_i/T_{2,CSF}} + \\
GM_{vol} * & \left[1 - e^{-Tl_i/T_{1,GM}} - (M_{GM}(t=0))e^{-Tl_i/T_{1,GM}} \right] e^{-TE_i/T_{2,GM}} + \\
WM_{vol} * & \left[1 - e^{-Tl_i/T_{1,WM}} - (M_{WM}(t=0))e^{-Tl_i/T_{1,GM}} \right] e^{-TE_i/T_{2,WM}} + \\
CBV * & \left[1 - e^{-Tl_i/T_{1,blood}} - (M_{blood}(t=0))e^{-Tl_i/T_{1,blood}} \right] e^{-TE_i/T_{2,blood}}
\end{aligned}$$

$$\begin{aligned}
S(Tl_i, TE_2, vol) = & \\
CSF_{vol} * & \left[1 - e^{-Tl_i/\tau_{1,CSF}} - (M_{CSF}(t=0))e^{-Tl_i/\tau_{1,CSF}} \right] e^{-TE_2/\tau_{2,CSF}} + \\
GM_{vol} * & \left[1 - e^{-Tl_i/\tau_{1,GM}} - (M_{GM}(t=0))e^{-Tl_i/\tau_{1,GM}} \right] e^{-TE_2/\tau_{2,GM}} + \\
WM_{vol} * & \left[1 - e^{-Tl_i/\tau_{1,WM}} - (M_{WM}(t=0))e^{-Tl_i/\tau_{1,GM}} \right] e^{-TE_2/\tau_{2,WM}} + \\
CBV * & \left[1 - e^{-Tl_i/\tau_{1,blood}} - (M_{blood}(t=0))e^{-Tl_i/\tau_{1,blood}} \right] e^{-TE_2/\tau_{2,blood}}
\end{aligned}$$

$$\begin{aligned}
S(TI_1, TE_3, vol) = & \\
CSF_{vol} * & \left[1 - e^{-T_{1,CSF}/T_1} - (M_{CSF}(t=0))e^{-T_{1,CSF}/T_1} \right] e^{-TE_3/T_{1,CSF}} + \\
GM_{vol} * & \left[1 - e^{-T_{1,GM}/T_1} - (M_{GM}(t=0))e^{-T_{1,GM}/T_1} \right] e^{-TE_3/T_{1,GM}} + \\
WM_{vol} * & \left[1 - e^{-T_{1,WM}/T_1} - (M_{WM}(t=0))e^{-T_{1,WM}/T_1} \right] e^{-TE_3/T_{1,WM}} + \\
CBV * & \left[1 - e^{-T_{1,blood}/T_1} - (M_{blood}(t=0))e^{-T_{1,blood}/T_1} \right] e^{-TE_3/T_{1,blood}}
\end{aligned}$$

$$\begin{aligned}
S(TI_2, TE_1, vol) = & \\
CSF_{vol} * & \left[1 - e^{-T_{I_2}/T_{1,CSF}} - (M_{CSF}(t=0))e^{-T_{I_2}/T_{1,CSF}} \right] e^{-TE_1/T_{2,CSF}} + \\
GM_{vol} * & \left[1 - e^{-T_{I_2}/T_{1,GM}} - (M_{GM}(t=0))e^{-T_{I_2}/T_{1,GM}} \right] e^{-TE_1/T_{2,GM}} + \\
WM_{vol} * & \left[1 - e^{-T_{I_2}/T_{1,WM}} - (M_{WM}(t=0))e^{-T_{I_2}/T_{1,GM}} \right] e^{-TE_1/T_{2,WM}} + \\
CBV * & \left[1 - e^{-T_{I_2}/T_{1,blood}} - (M_{blood}(t=0))e^{-T_{I_2}/T_{1,blood}} \right] e^{-TE_1/T_{2,blood}}
\end{aligned}$$

$$\begin{aligned}
S(Tl_2, TE_2, vol) = & \\
CSF_{vol} * & \left[1 - e^{-Tl_2/\tau_{1,CSF}} - (M_{CSF}(t=0))e^{-Tl_2/\tau_{1,CSF}} \right] e^{-TE_2/\tau_{2,CSF}} + \\
GM_{vol} * & \left[1 - e^{-Tl_2/\tau_{1,GM}} - (M_{GM}(t=0))e^{-Tl_2/\tau_{1,GM}} \right] e^{-TE_2/\tau_{2,GM}} + \\
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\end{aligned}$$

$$\begin{aligned}
S(Tl_1, TE_3, vol) = & \\
CSF_{vol} * & \left[1 - e^{-Tl_1/\tau_{1,CSF}} - (M_{CSF}(t=0))e^{-Tl_1/\tau_{1,CSF}} \right] e^{-TE_3/\tau_{2,CSF}} + \\
GM_{vol} * & \left[1 - e^{-Tl_1/\tau_{1,GM}} - (M_{GM}(t=0))e^{-Tl_1/\tau_{1,GM}} \right] e^{-TE_3/\tau_{2,GM}} + \\
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CBV * & \left[1 - e^{-Tl_1/\tau_{1,blood}} - (M_{blood}(t=0))e^{-Tl_1/\tau_{1,blood}} \right] e^{-TE_3/\tau_{2,blood}}
\end{aligned}$$

$$\begin{aligned}
S(Tl_3, TE_1, vol) = & \\
CSF_{vol} * & \left[1 - e^{-Tl_3/T_{1,CSF}} - (M_{CSF}(t=0))e^{-Tl_3/T_{1,CSF}} \right] e^{-TE_1/T_{2,CSF}} + \\
GM_{vol} * & \left[1 - e^{-Tl_3/T_{1,GM}} - (M_{GM}(t=0))e^{-Tl_3/T_{1,GM}} \right] e^{-TE_1/T_{2,GM}} + \\
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\end{aligned}$$

$$\begin{aligned}
S(Tl_3, TE_2, vol) = & \\
CSF_{vol} * & \left[1 - e^{-Tl_3/\tau_{1,CSF}} - (M_{CSF}(t=0))e^{-Tl_3/\tau_{1,CSF}} \right] e^{-TE_2/\tau_{2,CSF}} + \\
GM_{vol} * & \left[1 - e^{-Tl_3/\tau_{1,GM}} - (M_{GM}(t=0))e^{-Tl_3/\tau_{1,GM}} \right] e^{-TE_2/\tau_{2,GM}} + \\
WM_{vol} * & \left[1 - e^{-Tl_3/\tau_{1,WM}} - (M_{WM}(t=0))e^{-Tl_3/\tau_{1,GM}} \right] e^{-TE_2/\tau_{2,WM}} + \\
CBV * & \left[1 - e^{-Tl_3/\tau_{1,blood}} - (M_{blood}(t=0))e^{-Tl_3/\tau_{1,blood}} \right] e^{-TE_2/\tau_{2,blood}}
\end{aligned}$$

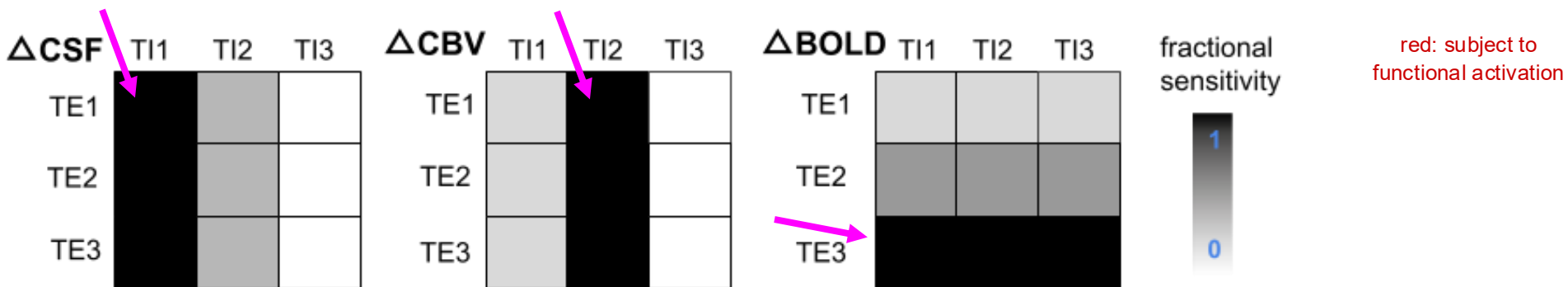
$$S(TI_3, TE_3, vol) =$$

$$CSF_{vol} * \left[1 - e^{-T_{I_3}/T_{I_3,CSF}} - (M_{CSF}(t=0)) e^{-T_{I_3}/T_{I_3,CSF}} \right] e^{-TE_3/T_{I_3,CSF}} +$$

$$GM_{vol} * \left[1 - e^{-T_{I_3}/T_{I_3,GM}} - (M_{GM}(t=0)) e^{-T_{I_3}/T_{I_3,GM}} \right] e^{-TE_3/T_{I_3,GM}} +$$

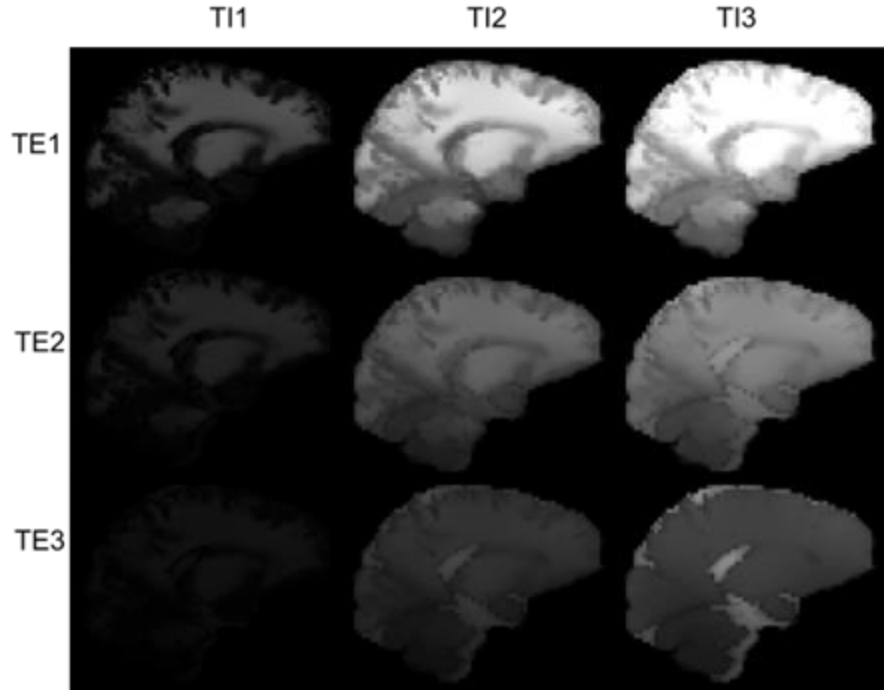
$$WM_{vol} * \left[1 - e^{-T_{I_3}/T_{I_3,WM}} - (M_{WM}(t=0)) e^{-T_{I_3}/T_{I_3,WM}} \right] e^{-TE_3/T_{I_3,WM}} +$$

$$CBV * \left[1 - e^{-T_{I_3}/T_{I_3,blood}} - (M_{blood}(t=0)) e^{-T_{I_3}/T_{I_3,blood}} \right] e^{-TE_3/T_{I_3,blood}}$$

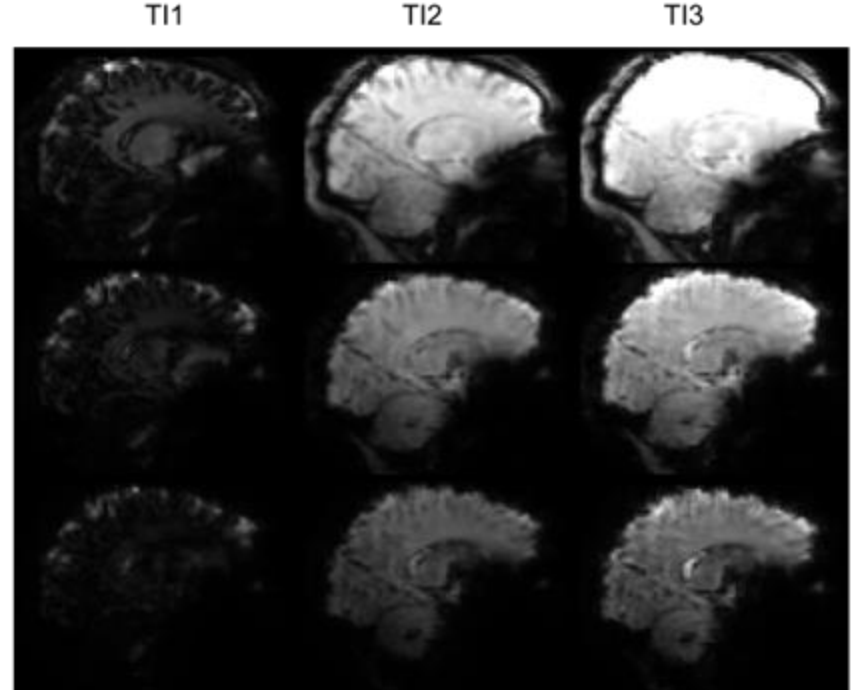


Confirming the method

Simulation

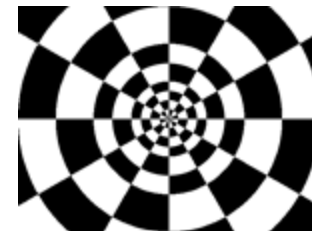
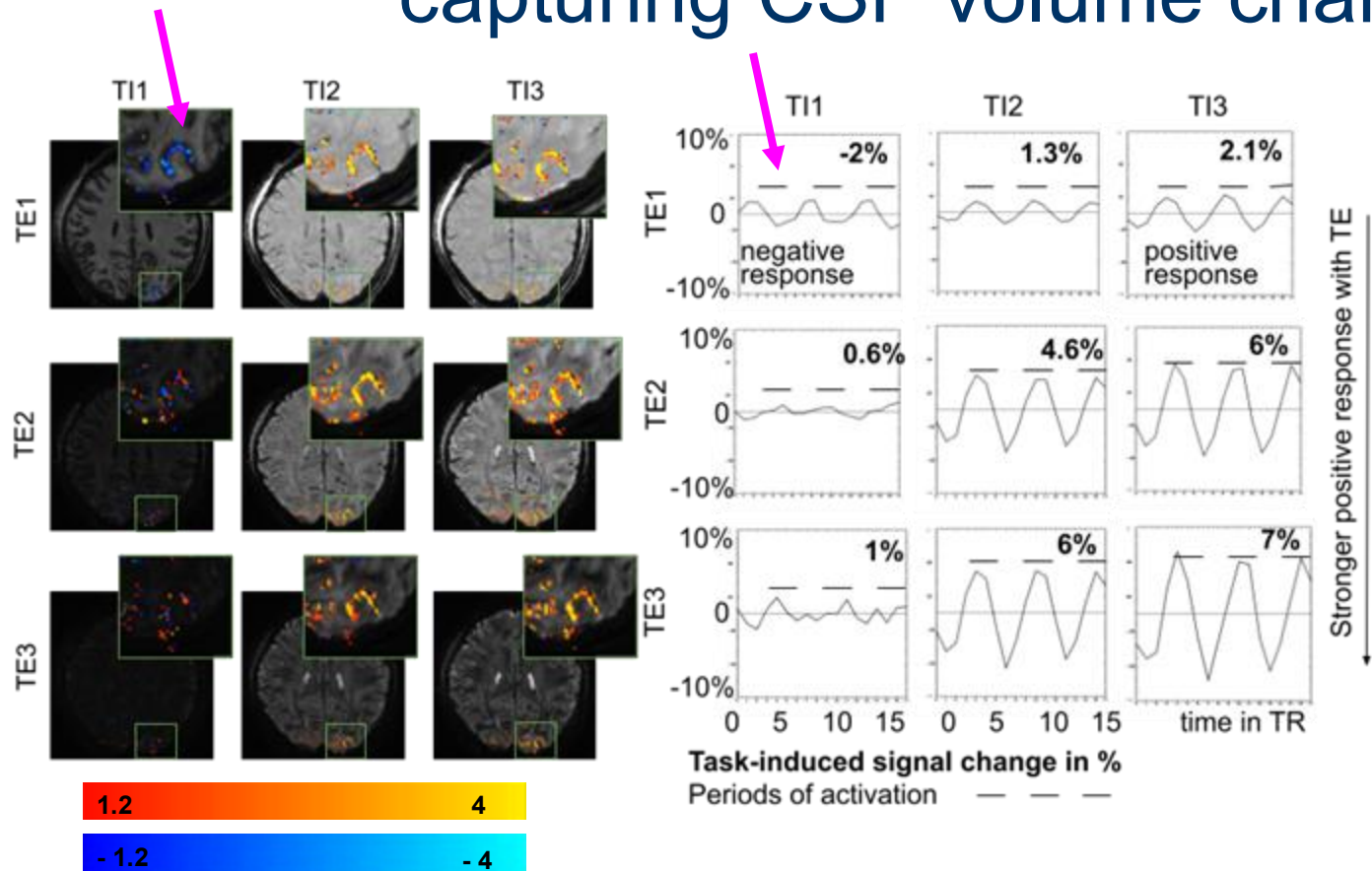


Empirical results



Comparing predicted tissue contrasts and the contrasts seen in the empirical data shows that overall our predictions match the empirical results

Task-locked fluctuations indicate the sequence capturing CSF volume change

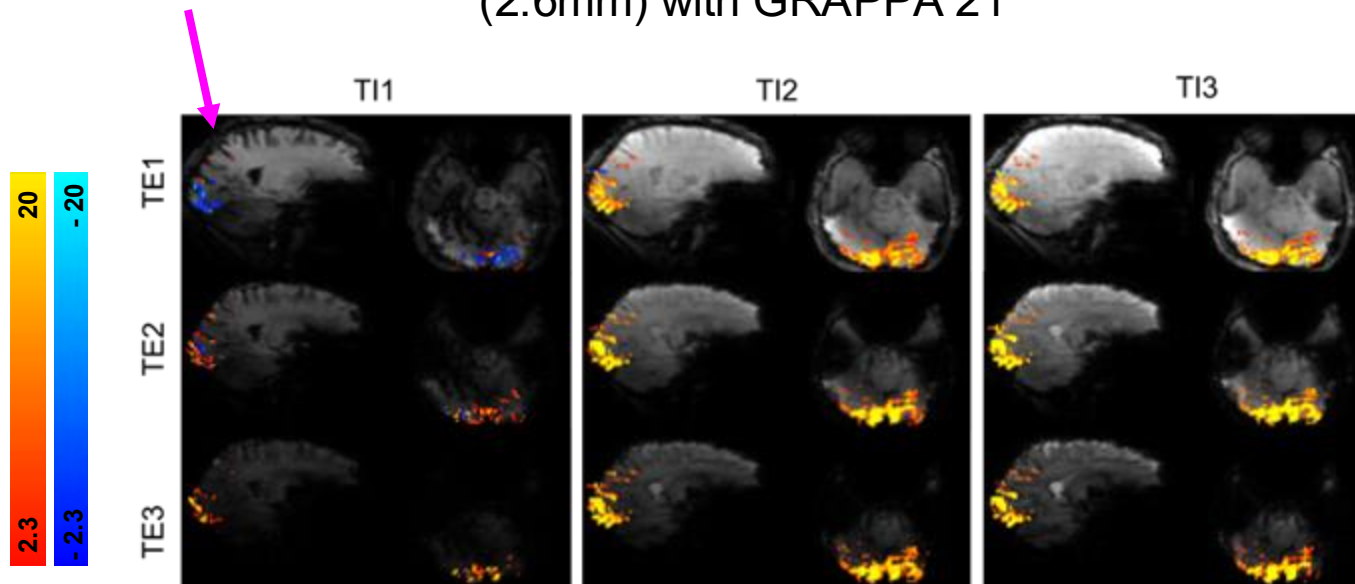
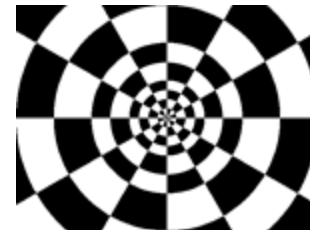


Negative signal changes seen with the early TI, while later TIs resulted in increasingly positive signal changes

Functional responses are stronger at later echo times, as expected from GE-BOLD

Task-locked fluctuations indicate the sequence capturing CSF volume change

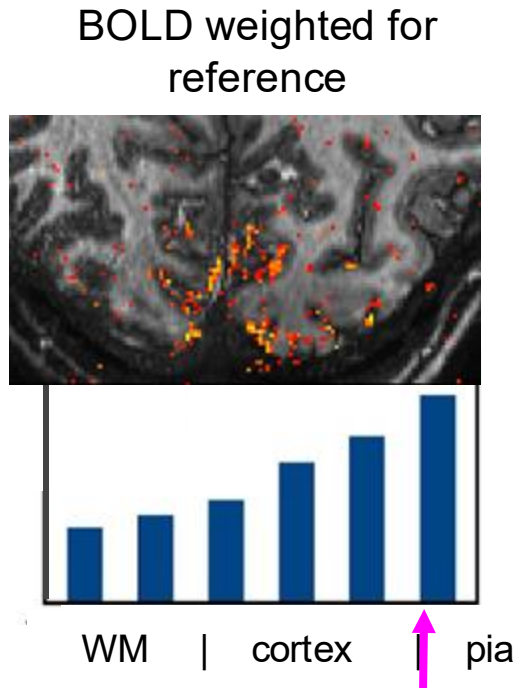
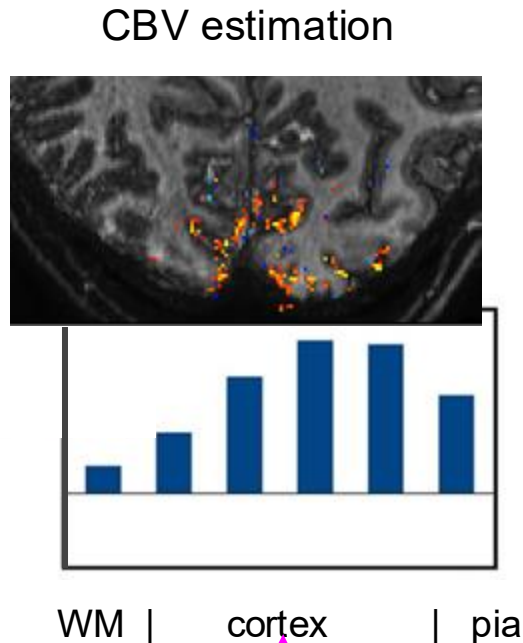
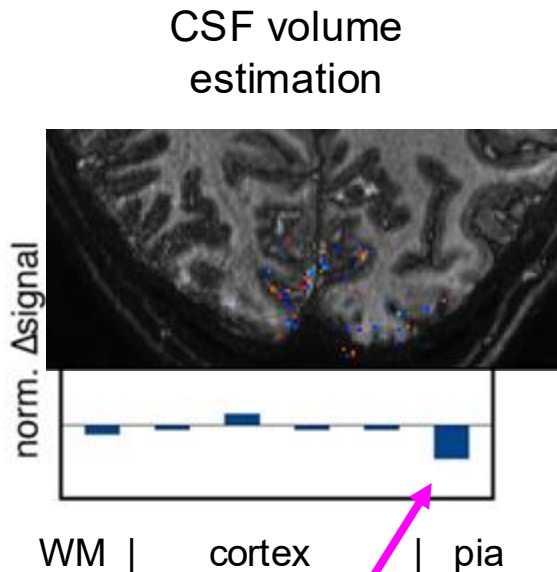
Feasibility of 3D whole brain mapping at low resolution (2.6mm) with GRAPPA 21



Task-induced activation in the visual cortex can be seen at high resolution, in time courses and at whole brain resolution

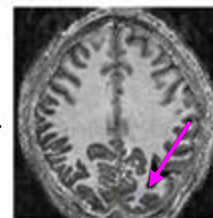
Confirmation that this method can spatially map CSF

functional responses,
tissue type volume
estimates



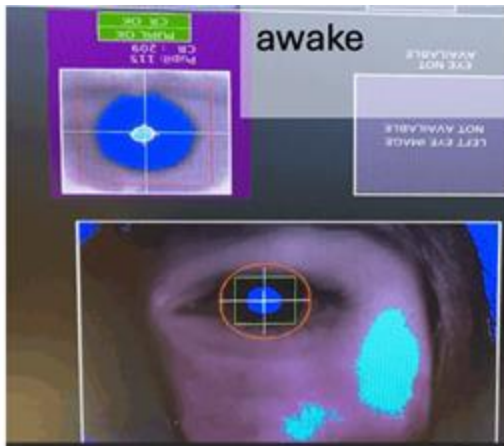
- Indications of a small CSF volume reduction in voxels above the cortical ribbon, consistent with previous work (Jin, 2010)
- CSF-weighted signal changes are largest above the cortex

ROI for layer extraction



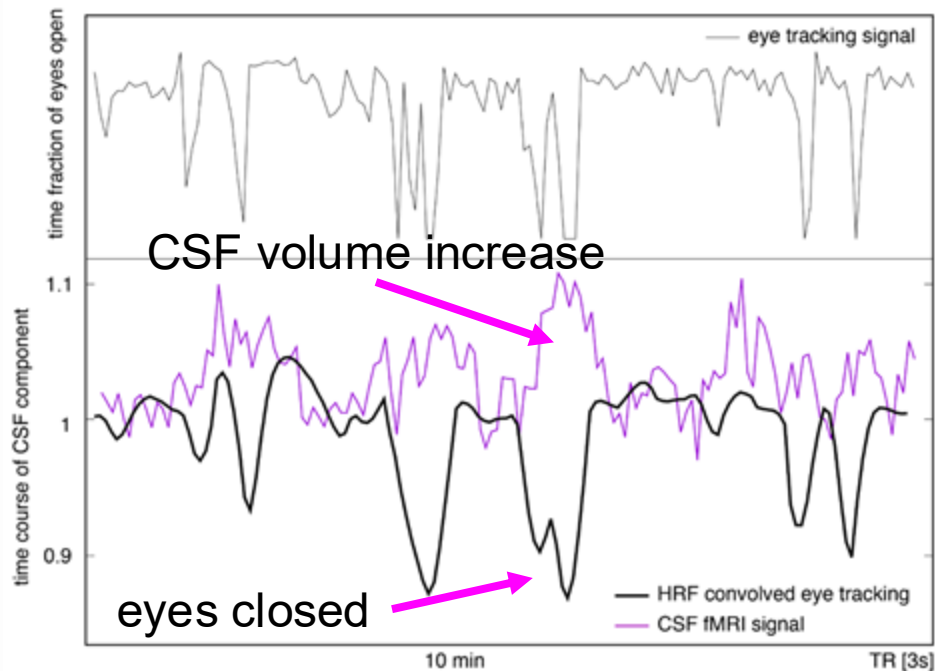
CSF volume increases as participants get drowsy

Tacking vigilance: Eye tracker for pupil measurement in the scanner room



Eye tracking system shown on experimenter

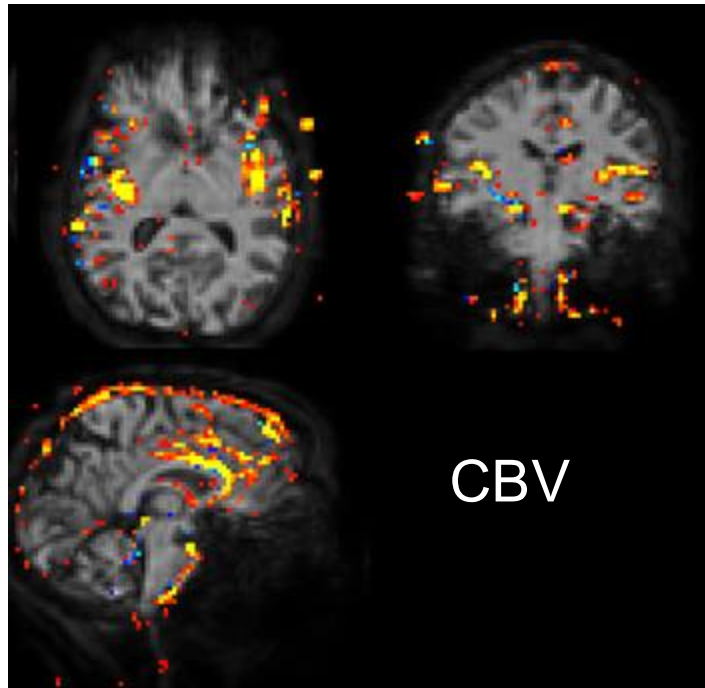
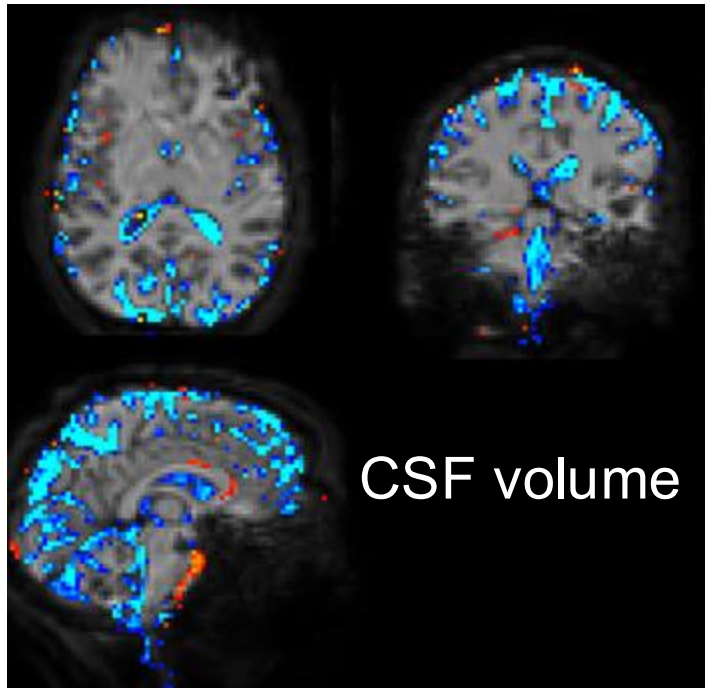
Time course of “sleepy” eyes vs. CSF volume
(picked one section of three runs)



As eyes get more drowsy, CSF volume increases

CSF and CBV correlates of alertness

Preliminary maps of CSF and CBV correlates of alertness



CSF shows a decrease with alertness and CBV shows an increase with alertness

Summary and Conclusion

- We developed a multi-echo multi-inversion fMRI sequence that is simultaneously sensitive to CSF volume, CBV and BOLD.
- We see indications of CSF volume change during functional activation. And even more so during changes of drowsiness.
- Future work will include further validations of the Bloch model (converting 9 TI-TE combinations $\rightarrow$ CSF, CBV, BOLD) with ground truth CSF modulation tasks.

Thank you!

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MRI scanning was performed in the FMRIF core.



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