

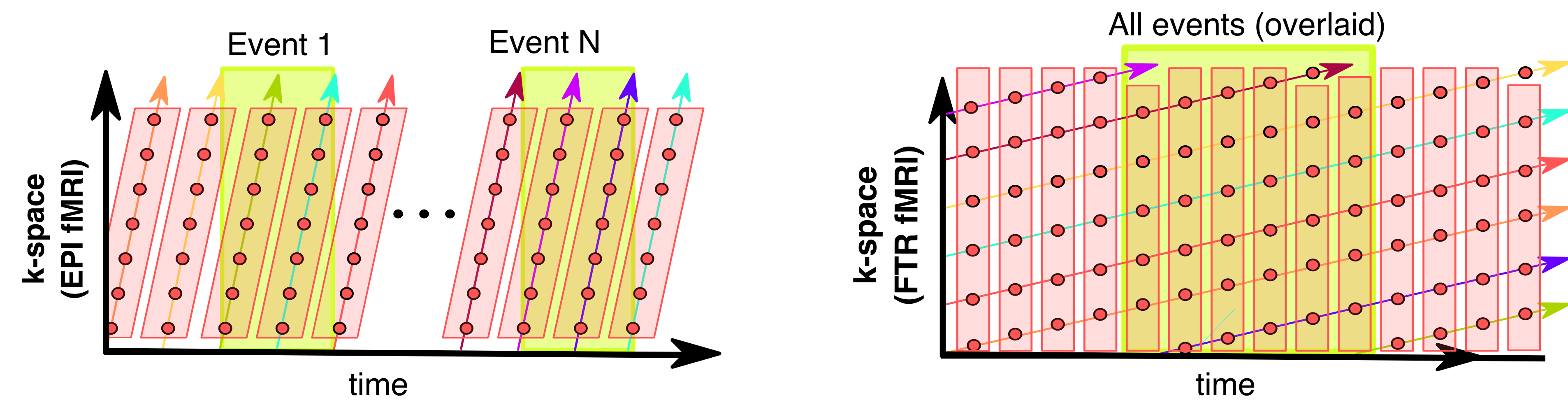
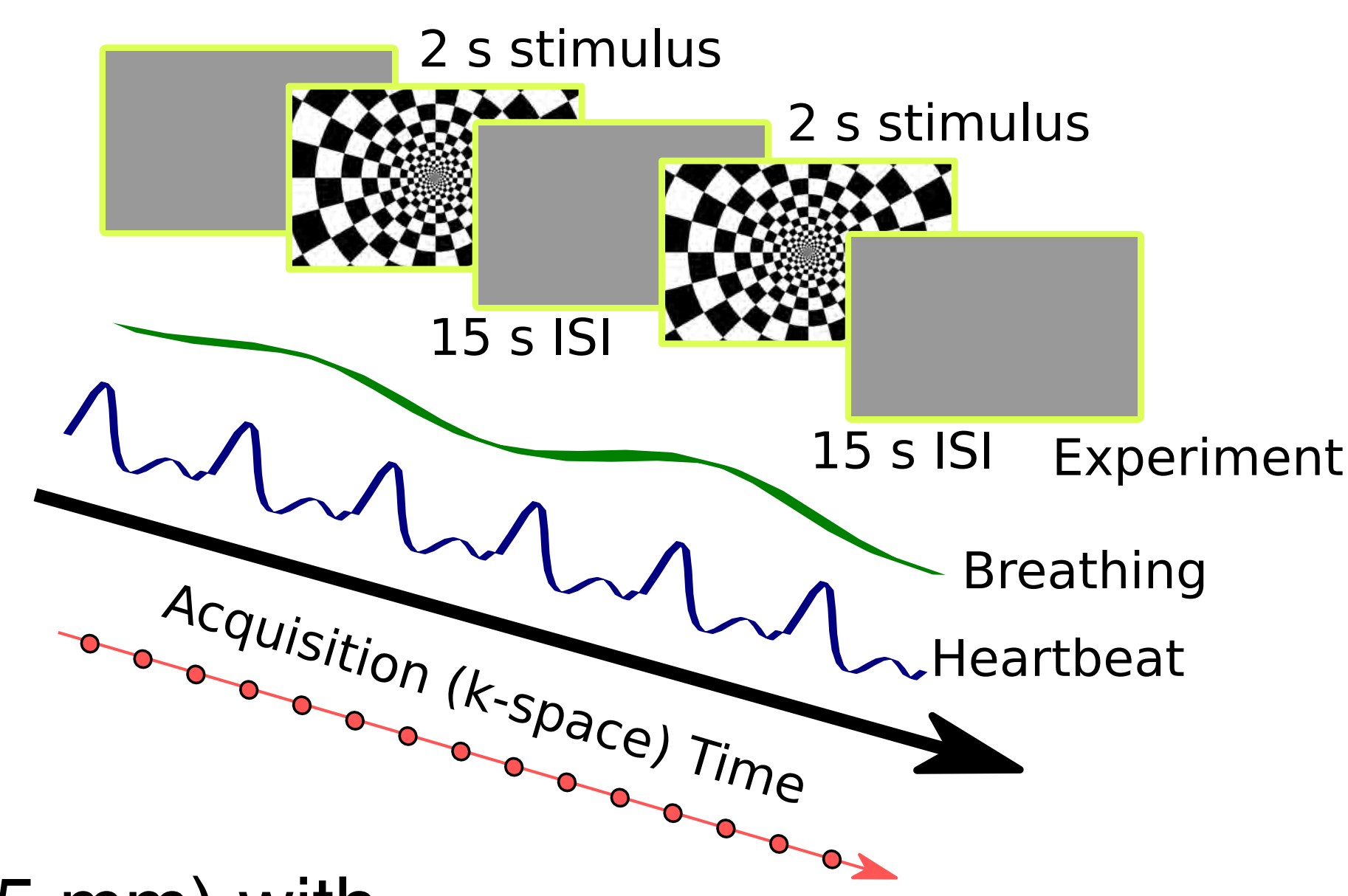
# A functionally time-resolved reconstruction technique for high-resolution fMRI.

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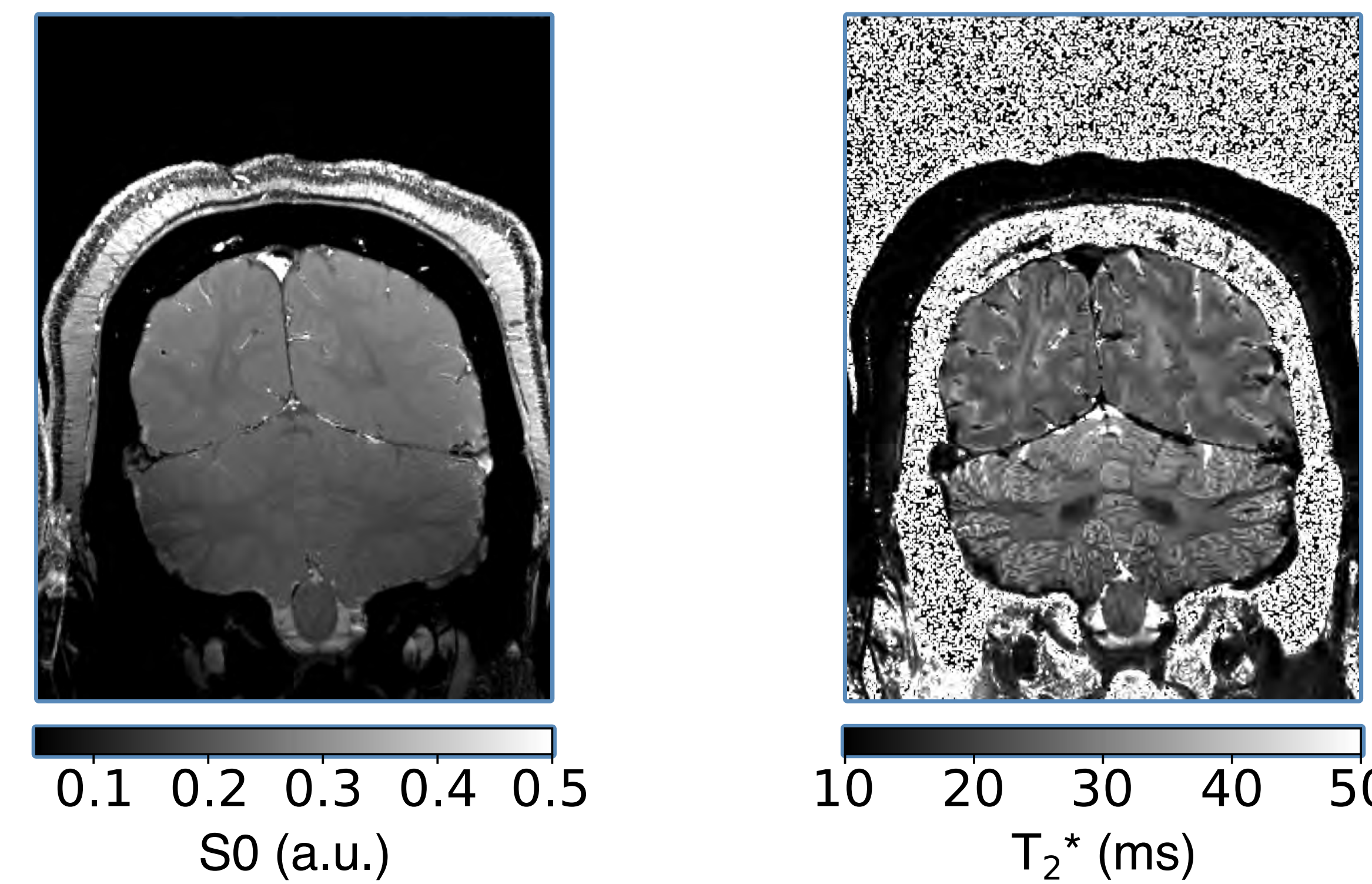
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## Introduction

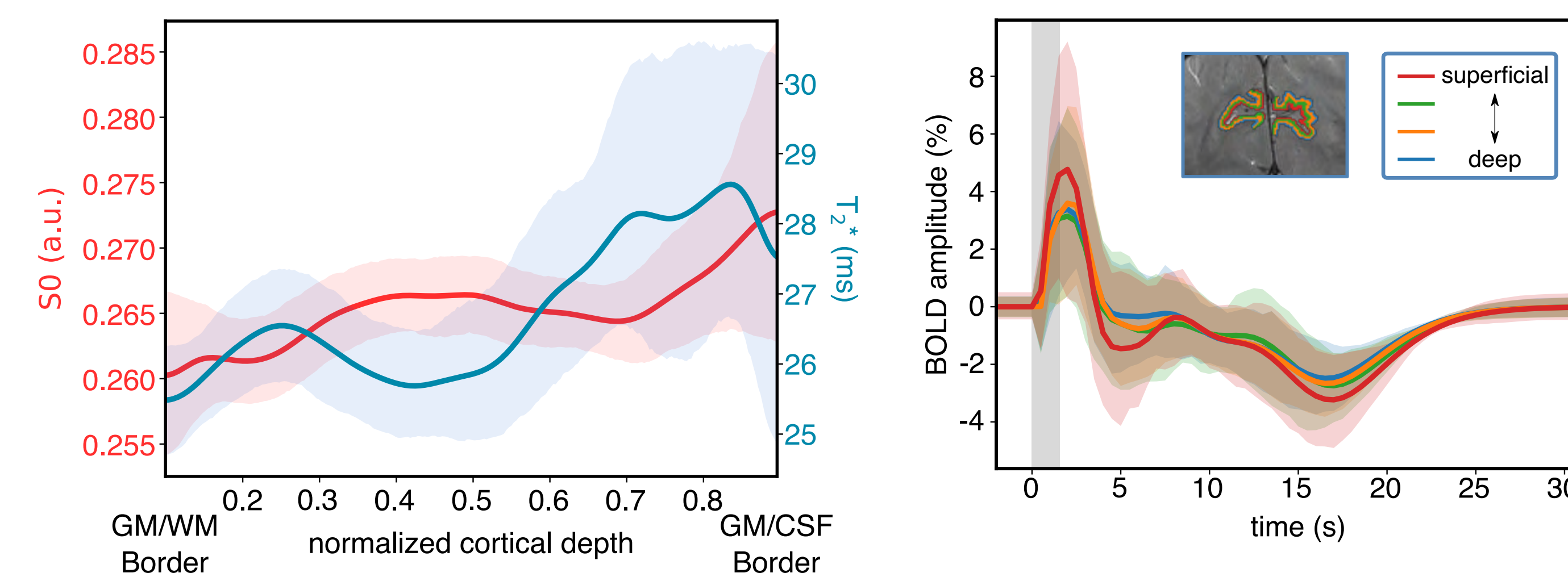
- High resolution fMRI is a promising method to probe mesoscopic brain responses<sup>1,2</sup>, yet is currently too coarse to sample from individual human columns and layers.
- High-resolution fMRI using EPI readouts and BOLD contrast suffer from spatial distortions and  $T_2^*$  blurring due to long readout trains<sup>3</sup>.
- Here, we build on time resolved methods<sup>4,5</sup> to incorporate neuroscientific experimental designs and physiological confounds into fMRI reconstruction to time resolve data from multi-echo, multi-shot gradient echo sequences (one k-space line to minimize readout time).
- Our method, **functionally time-resolved fMRI** (fTR), of very high spatial resolution (here, up to 0.5 mm) with multiple (6) echoes without sacrificing temporal resolution (here, 0.5 s).



## Results (Task Mapping)



**Reconstructed images.** Multi-echo data are used to fit high-resolution (0.5 mm)  $S_0$  and  $T_2^*$  decay time parameters. **Left:**  $S_0$  images display good gray/white matter and CSF contrast, enabling segmentation of functional data. **Right:**  $T_2^*$  images provide additional CSF and vascular contrast, as well as identification of striate cortex based on the visible Stria of Gennari.

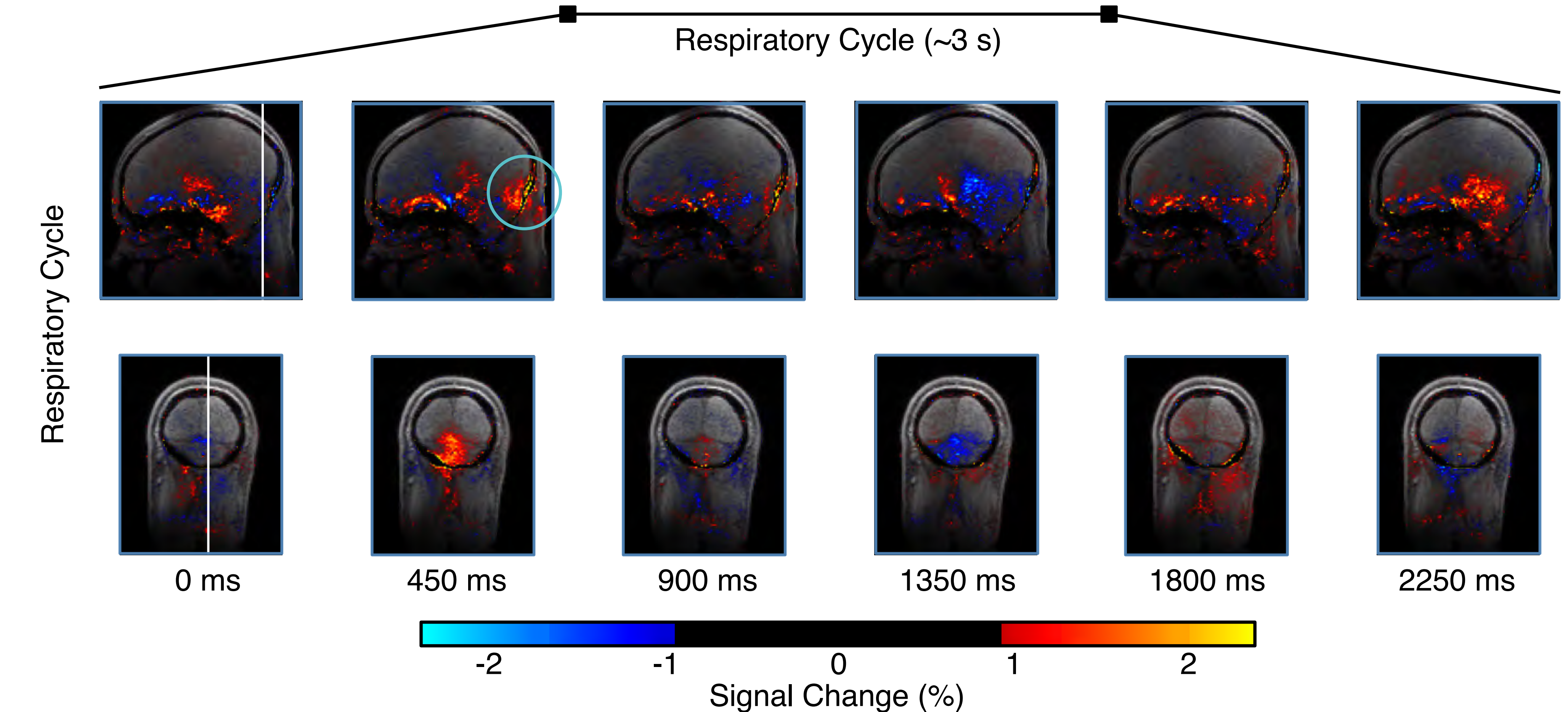
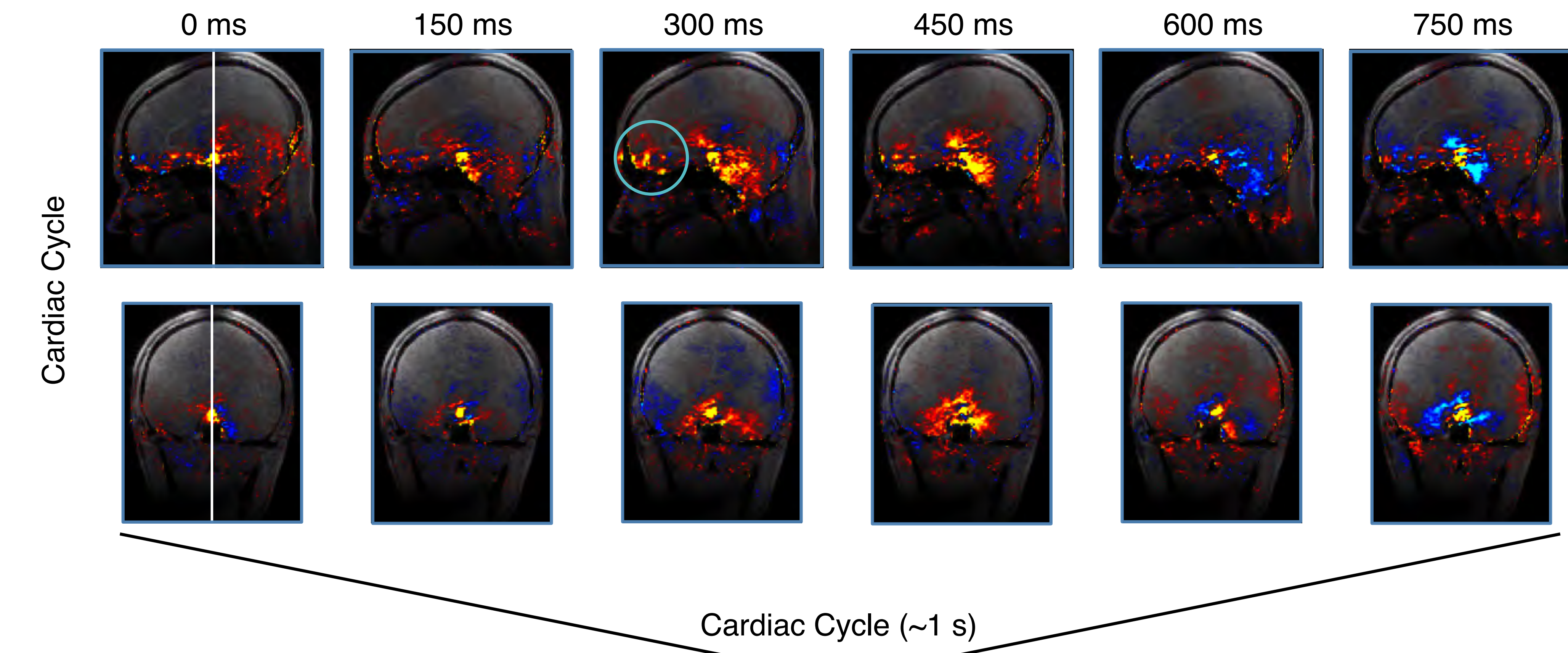


**Laminar analyses.** **Left:** Anatomical  $S_0$  and  $T_2^*$  values are plotted against normalized cortical depth values in the calcarine sulcus (see segmentation image on right). Shaded areas indicate 95% CIs on the mean laminar profile.  $T_2^*$  display a characteristic dip in middle layers of striate cortex. **Right:** Functional BOLD responses (optimally combined echo data) to a 2-second flashing checkerboard stimulus (10 Hz) are shown for 4 cortical depth ROIs (see inset segmentation). The gray area indicates stimulation period and shaded colors indicate 95% CIs on individual voxel time courses. Superficial layers (red) show higher response amplitudes and larger undershoots than other cortical depths, similar to high-resolution temporal BOLD profiles seen in rodents<sup>6</sup>.

## Conclusions

- Functionally time-resolved fMRI incorporates experimental designs and physiology into image reconstruction to capture high spatial and temporal resolution brain responses.
- fTR-fMRI does not suffer from spatial distortions in the phase encoding direction and  $S_0/T_2^*$  maps provide good anatomical contrast, aiding neuroscientific interpretation of experimental data.
- Future directions include combining task and physio mapping to better isolate task-related effects from layer fMRI responses.

## Results (Physiology Mapping)



**Reconstructed physiological cycles.** Whole brain MRI data (2 mm isotropic resolution) are resolved over physiological processes. Maps are thresholded to signal changes with magnitudes greater than 1%. **Upper:** A time course of one cardiac cycle (~1 s) is shown. Signal changes are primarily in subcortical areas, as previously reported<sup>9</sup>. The cyan circle highlights additional positive signal changes to the frontal pole during the late systole period. **Lower:** A time course of one respiratory cycle (~3 s). Signal change maps follow susceptibility changes above the frontal sinus; the cyan circle highlights additional changes in occipital cortex and the cerebellum, potentially due to breathing movement.

## References

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This work was supported by the NIH Intramural Program (#ZIAMH002783) and utilized computational resources from the NIH HPC Biowulf Cluster.

## Methods

- Acquisition:** Five subjects' data were collected on a Siemens MAGNETOM 7T+ w/ Nova 32Rx head coil. We collected two different sequences to demonstrate the utility of fTR-MRI: 1) a 2D GRE sequence (TR=31 ms, TEs=[4.22, 8.38, 12.54, 16.7, 20.86, 25.02] ms (bipolar readout), res=0.5x0.5 mm, slc thickness=0.8 mm, matrix=360x270, PE=R/L, repetitions=36, no acceleration or Partial Fourier, acquisition time=5:01). Prescription was perpendicular to the Calcarine Sulcus. 2) a whole brain 3D GRE sequence (TR=4.5 ms, TE=2.1 ms, res=2.0 mm isotropic, FOV=216x220x88 mm, PE=A/P, repetitions=70).
- Reconstruction:** Data were reconstructed via low-rank tensor completion<sup>4,7</sup> with modes for k-space, receivers, echoes and response time. Initial k-space, channel and echo subspaces were derived non-time resolved data with SVD. The time subspace was either an informed 2-gamma HRF basis (with derivatives; experimental mapping), or an order-2 Fourier basis (physio mapping). The subspaces and rank-(20, 7, 3, 5) core tensor were then iteratively updated with available data until the 2-norm of the core converged.
- Experiment:** Participants were presented a flashing radial checkerboard (10 Hz; 2 s presentation, 15 s ISI) and were asked to fixate for the entire experiment.
- Analysis:**  $S_0$  &  $T_2^*$  maps were linearly fit to log-multi-echo data. These maps were used to segment gray matter of the Calcarine sulcus region and to compute cortical depth values for each voxel (to plot layerwise  $S_0/T_2^*$  values) and 4 equivolume layers (to plot response functions) using LAYNI software<sup>8</sup>. 95% confidence intervals for  $S_0/T_2^*$  values were computed via 1000 bootstrap samples (33% of voxels in each sample) of layer profiles. CIs for response functions represent the 95% bounds of individual voxel responses.

