

The Magical World of **fMRIprep**

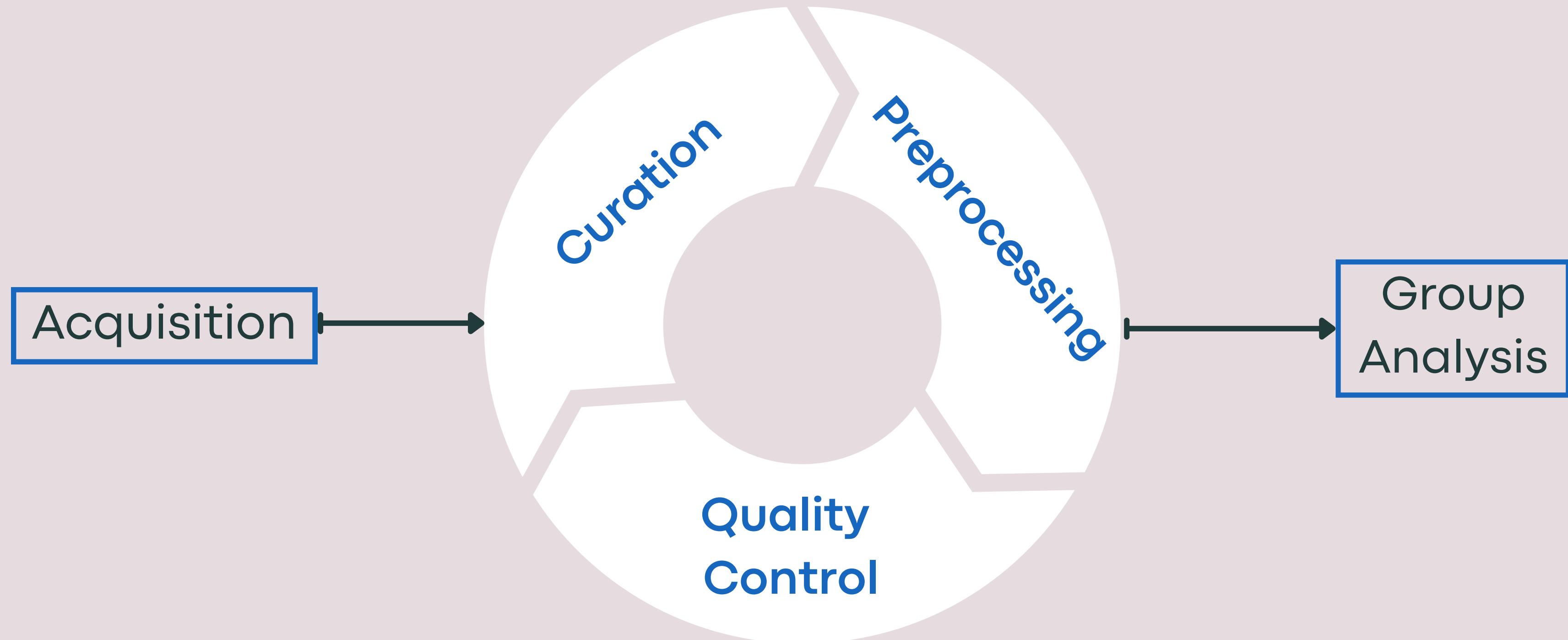
Rotation Recap by Margaret Gardner

Hi

My Objective:

Learn best-practice informatics
procedures for
reproducible & transparent
curation & analysis of fMRI data

fMRI Lifecycle



Day 2 Reward

n=125 subjects

anat = 125 subjects

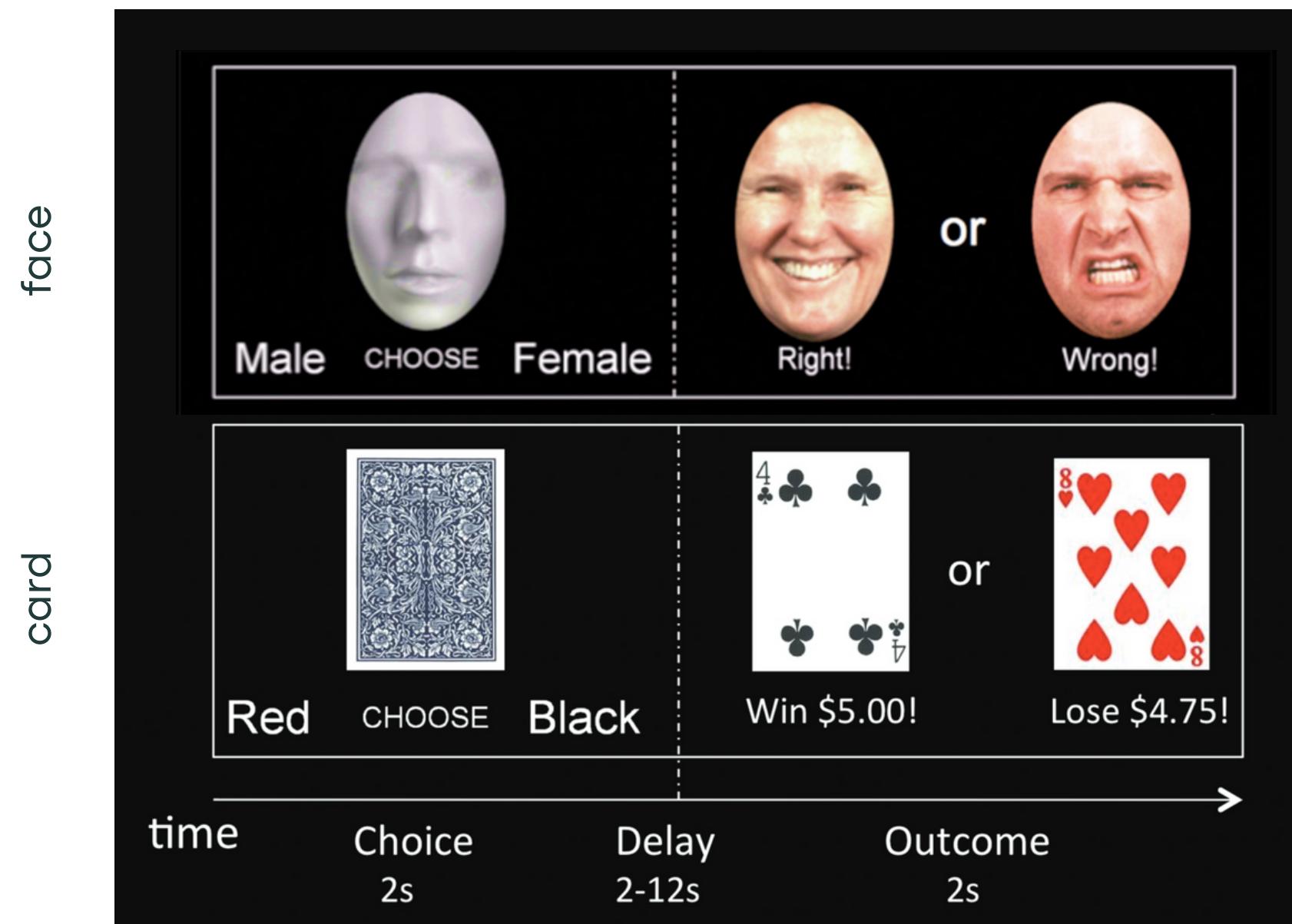
- T1w = 125
- T2w = 3

func = 125 subjects

- card run 01 = 124
- card run 02 = 124
- face run 01 = 123
- face run 02 = 124
- rest = 114

fmap = 124 subjects

dwi = 3 subjects



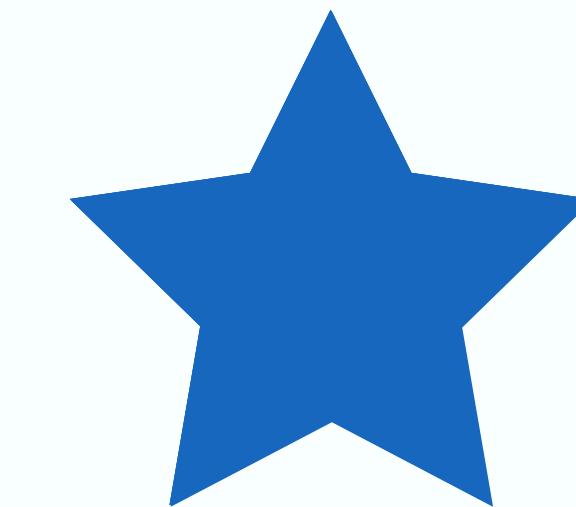
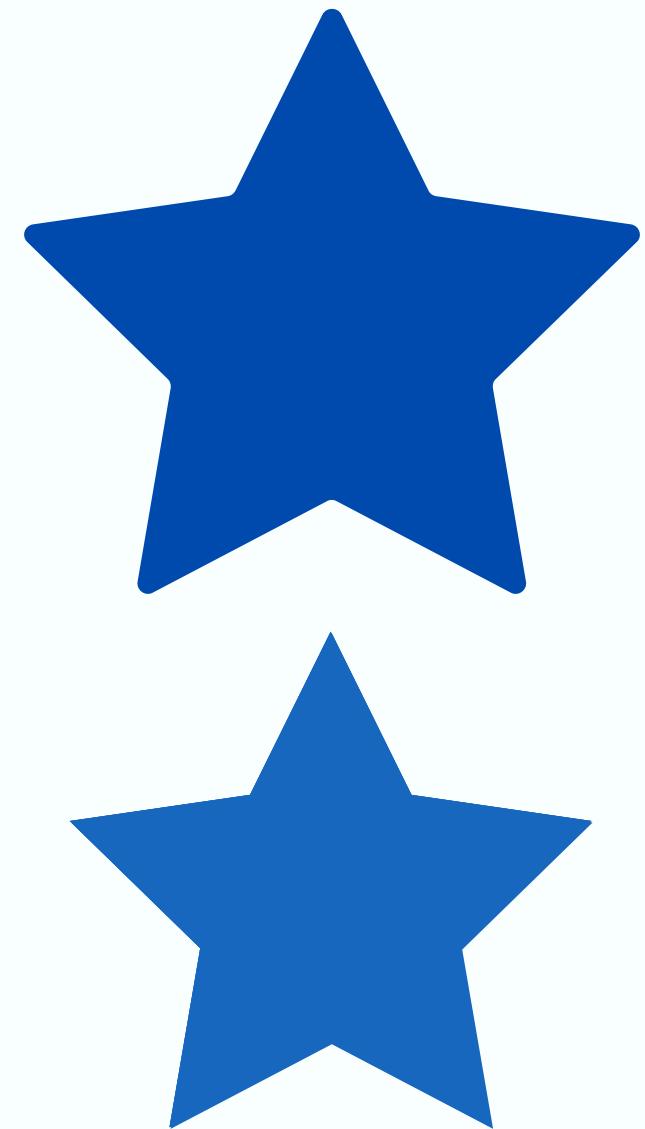
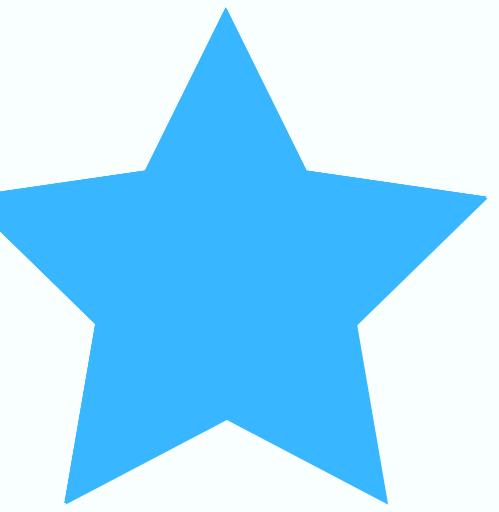
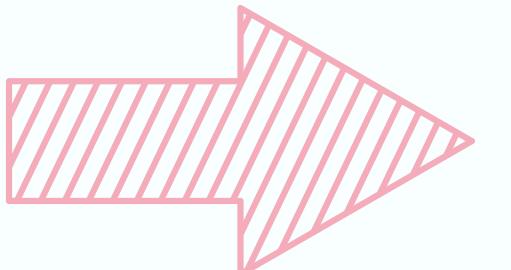
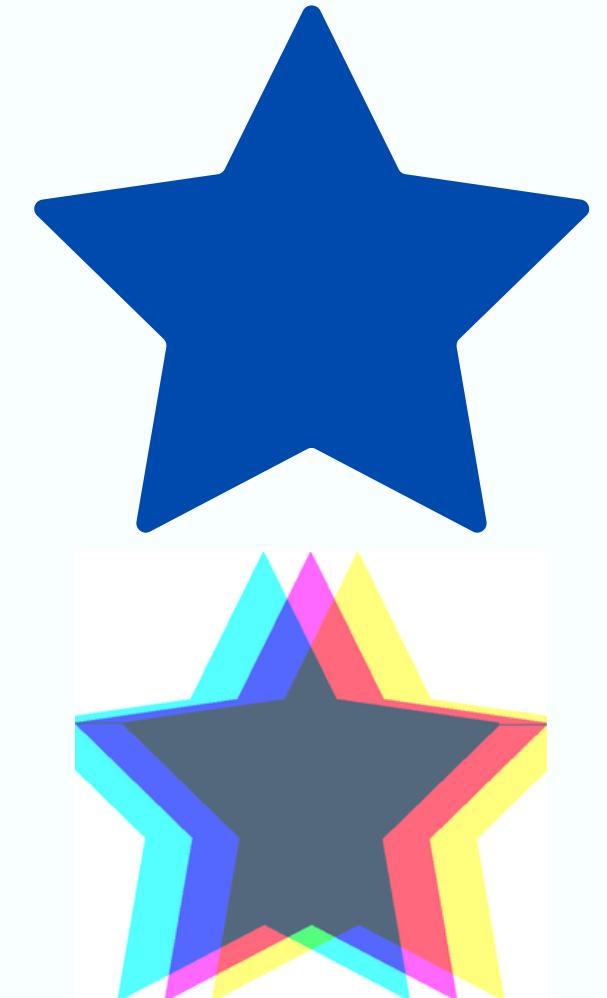
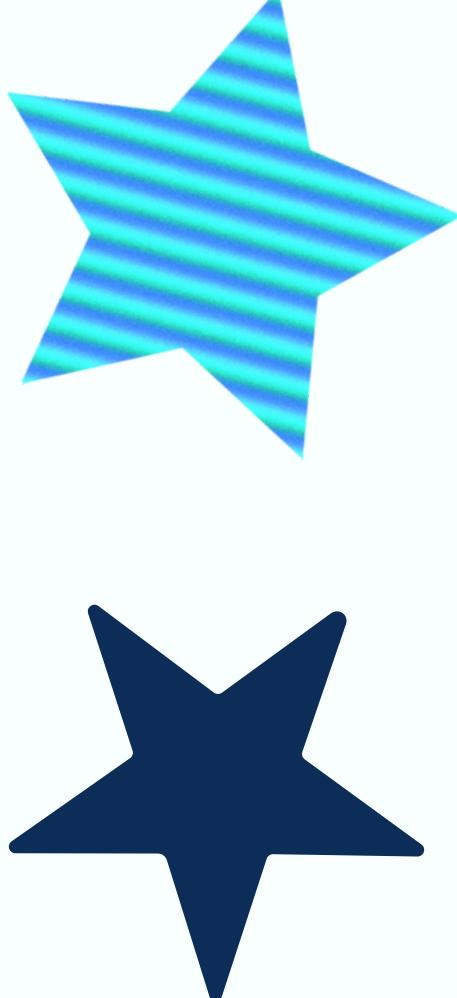
Adapted from Sharma et al, 2016 and Wolf et al, 2014

And now,

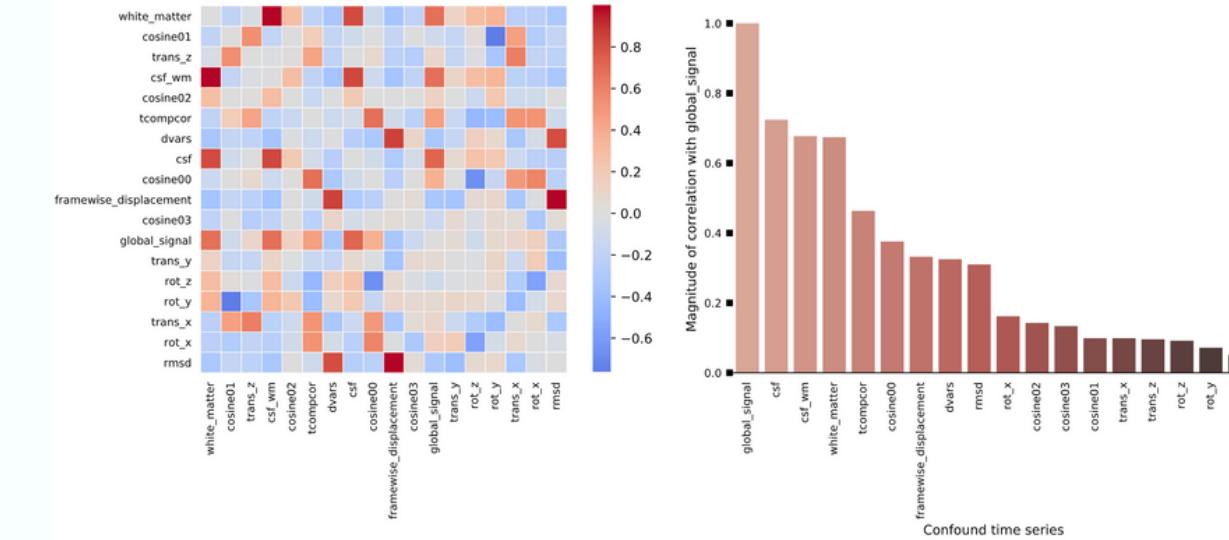
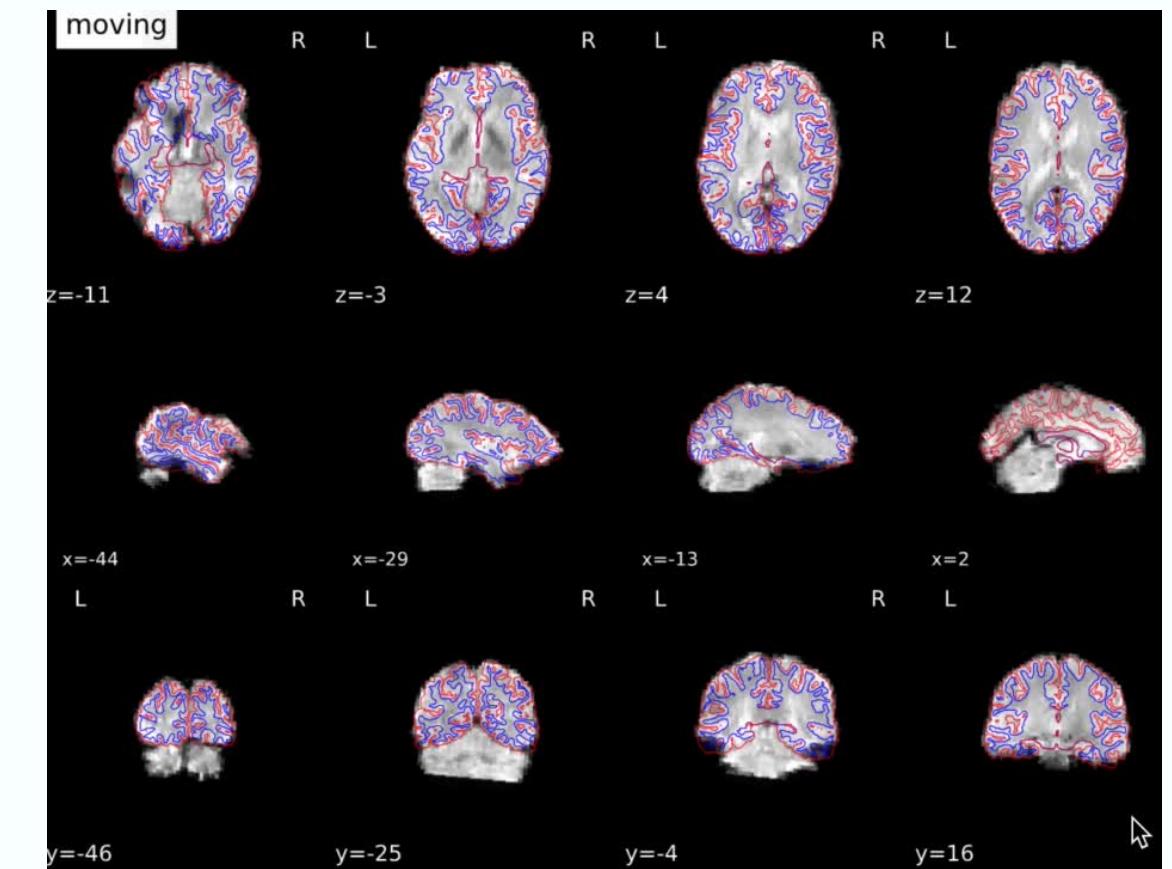
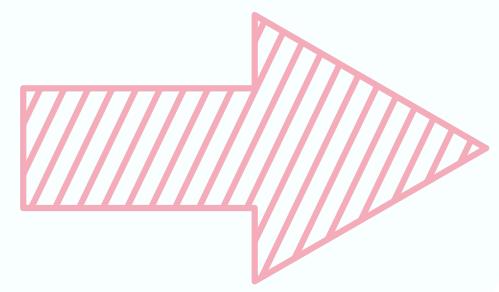
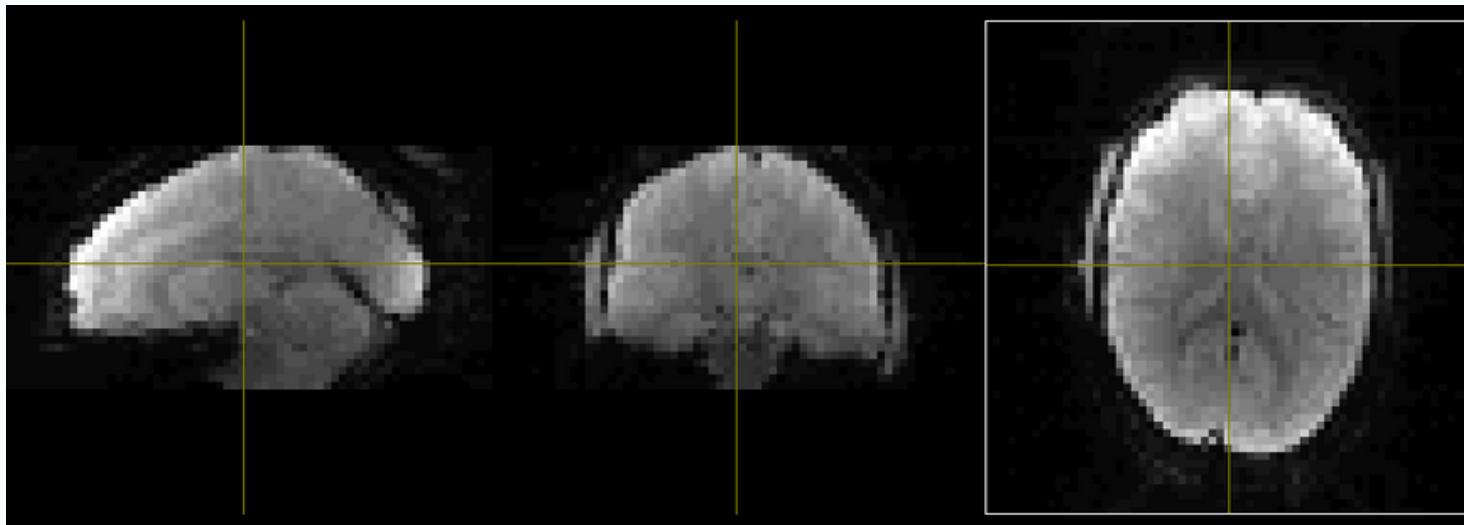
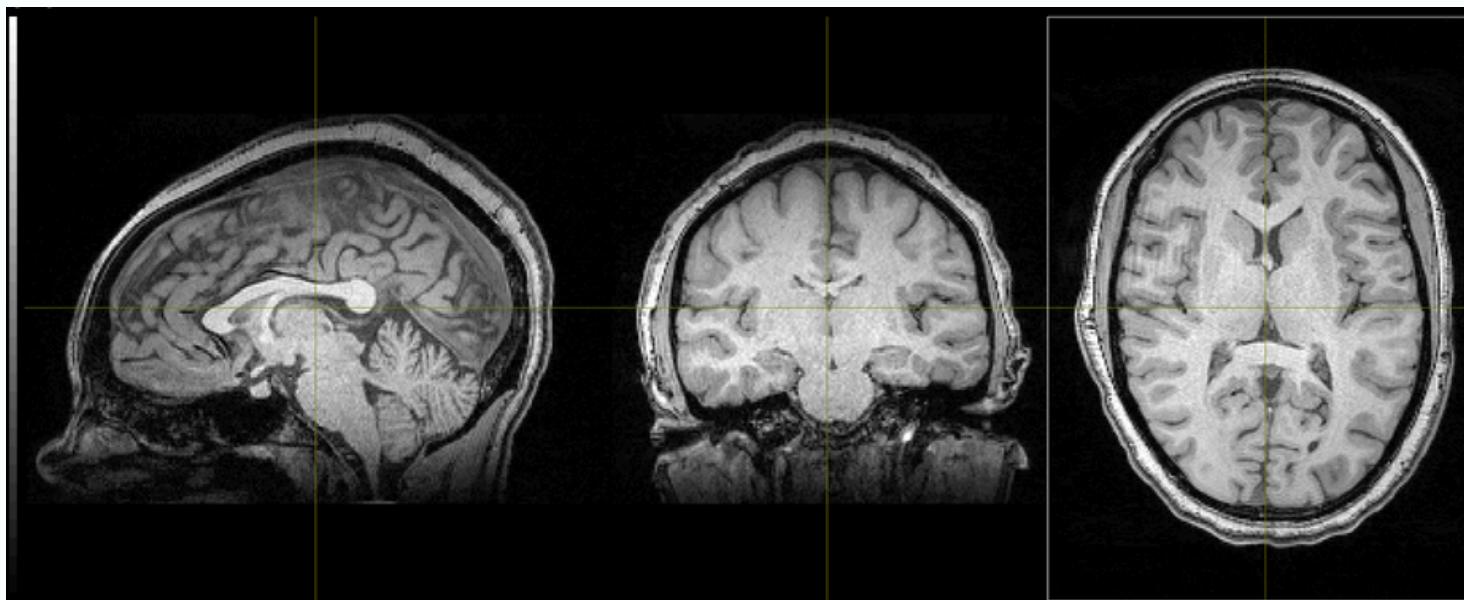
fMRIprep

What is preprocessing?

Cleaning up our data to more easily identify meaningful signal.



What is preprocessing?



Why fMRIprep?

AFNI: environment for processing and displaying fMRI data

ANTs: set of tools to normalize data to standard spaces

FSL: popular set of tools for analyzing many types of MR data

FreeSurfer: program for analysis of (mostly) structural images, especially for surface analyses

tedana: Python library of tools for denoising multi-echo fMRI

It's a BIDS App!

- Can intelligently read info from BIDS inputs
- Can automatically adapt processing based on inputs
- Runs on Docker/Singularity, so it's all the subprograms (packages) are self-contained and reproducible

It uses NiPype!

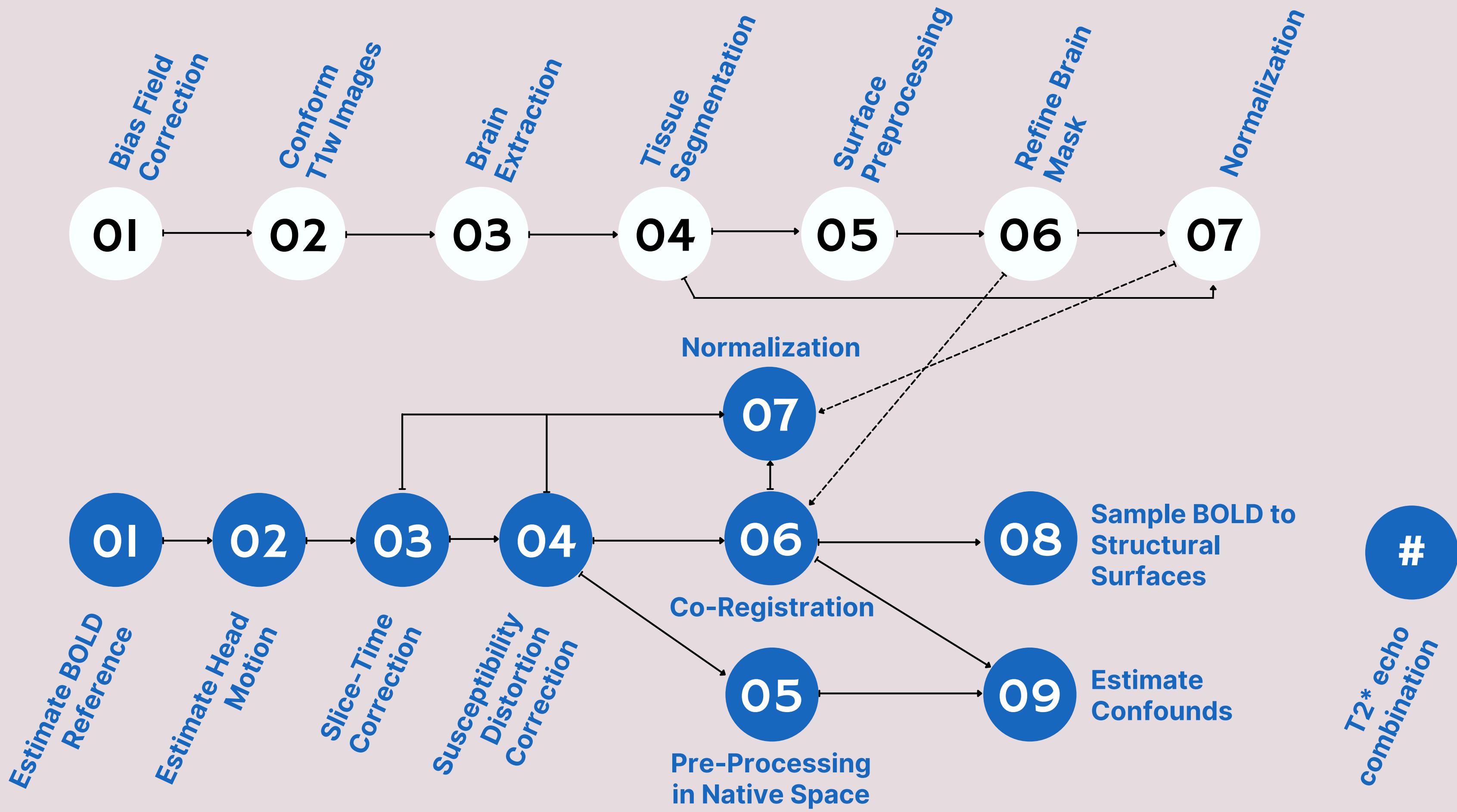
- Integrates scripts from different packages with Python

It gives you lots of options for subsequent processing!

- No spatial smoothing
- Calculates lots of confounds

It's transparent!

- Provides comprehensive visual QC reports automatically
- Boilerplate citation for manuscripts



*Bias Field
Correction*

01

*Conform
 $T1w$ Images*

02

*Brain
Extraction*

03

*Tissue
Segmentation*

04

*Surface
Preprocessing*

05

*Refine Brain
Mask*

06

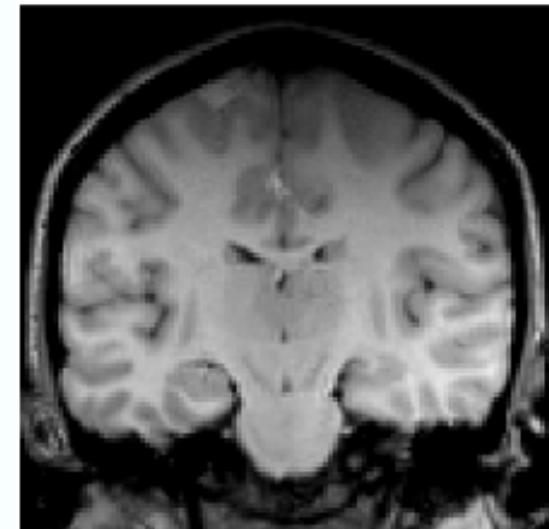
Normalization

07

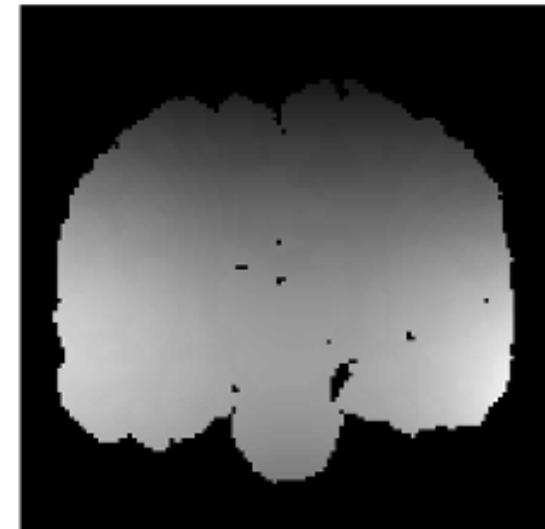
Structural Workflow

Bias Field Correction

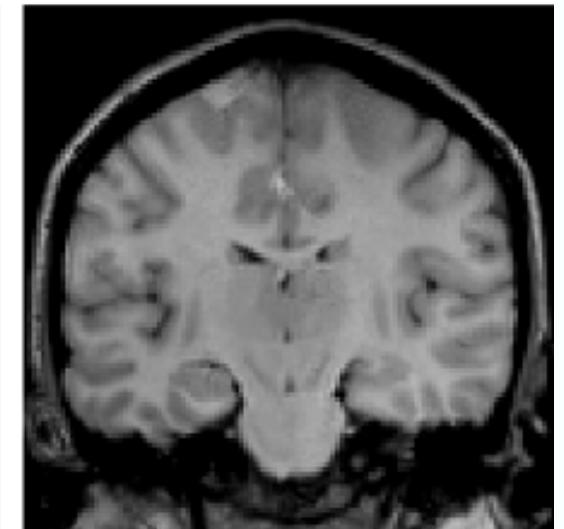
Original image



Inhomogeneity field



Corrected image



Vovk U, Pernus F & Likar B, 2007

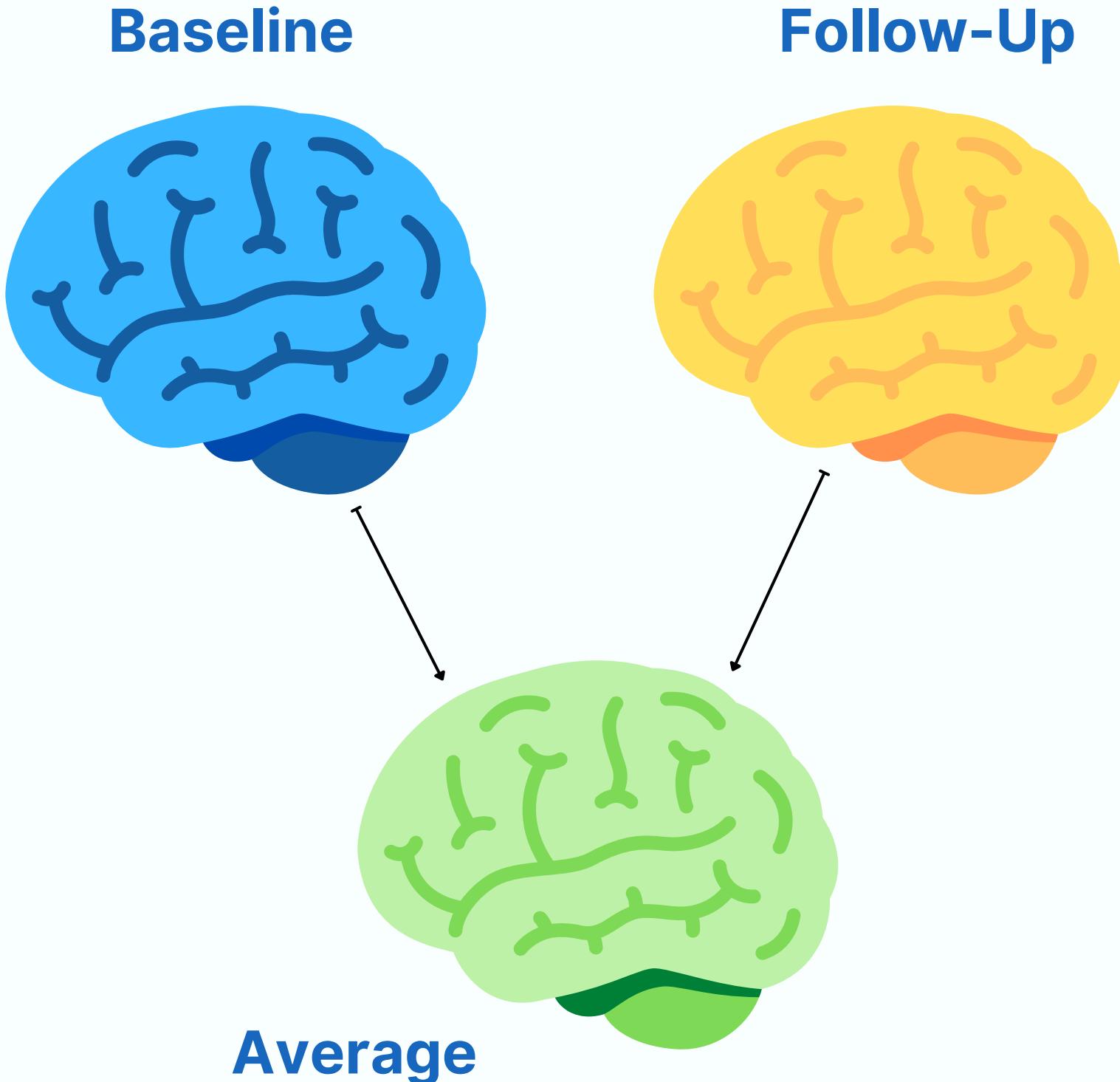
Identifies and corrects changes in voxel intensity caused by inhomogeneities in the magnetic field

- Uses *N4BiasFieldCorrection* from ANTs

Conform Tlw Images

Optional step to create a within-subject template for longitudinal studies in which you don't expect structural changes

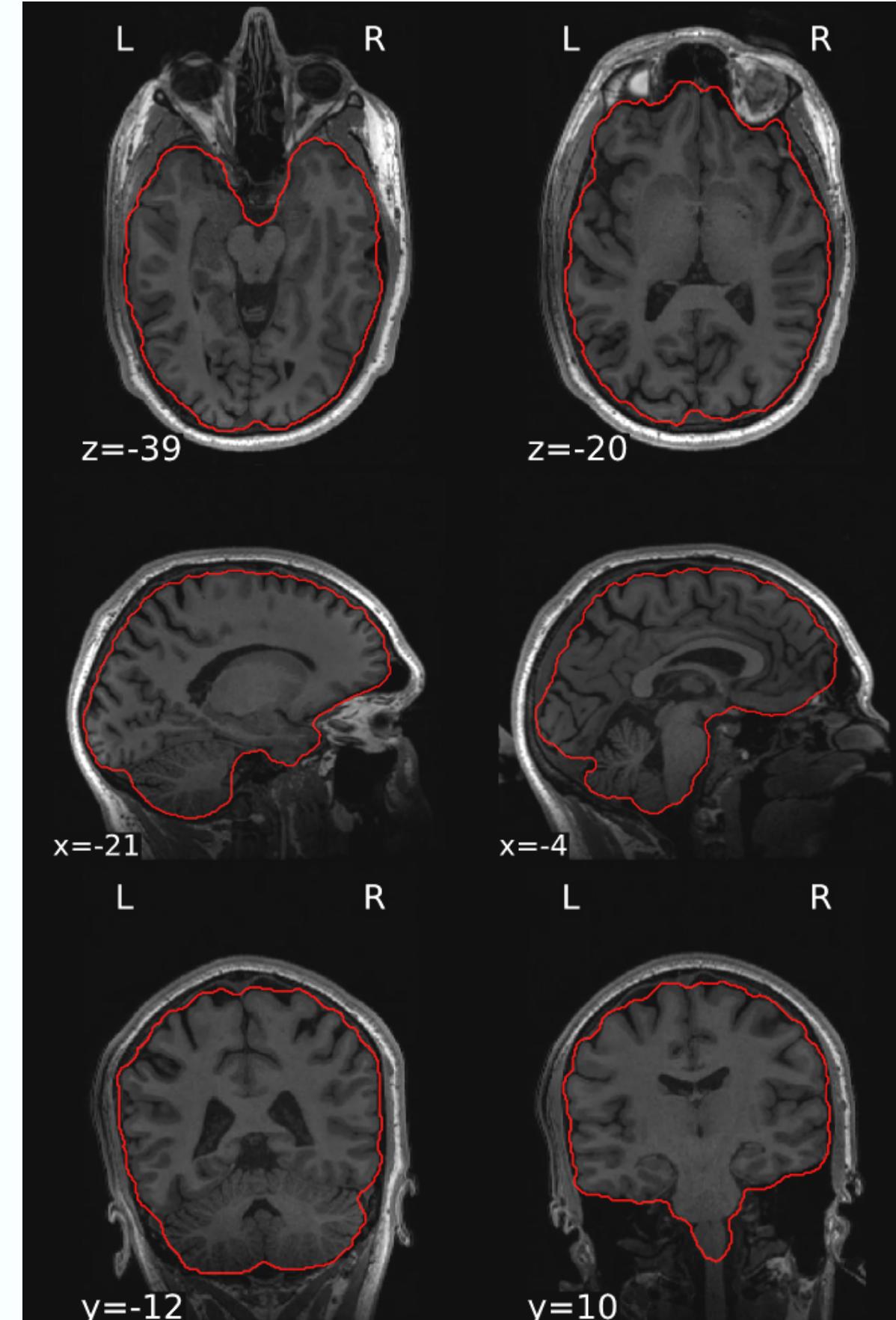
- Uses *mri_robust_template* from FreeSurfer
- Can be helpful (i.e. more evenly distributing the transforms needed to normalize baseline and follow-up)
- Unbiased (all images averaged) or aligned to first image



Brain Extraction

Removes all non-brain tissue

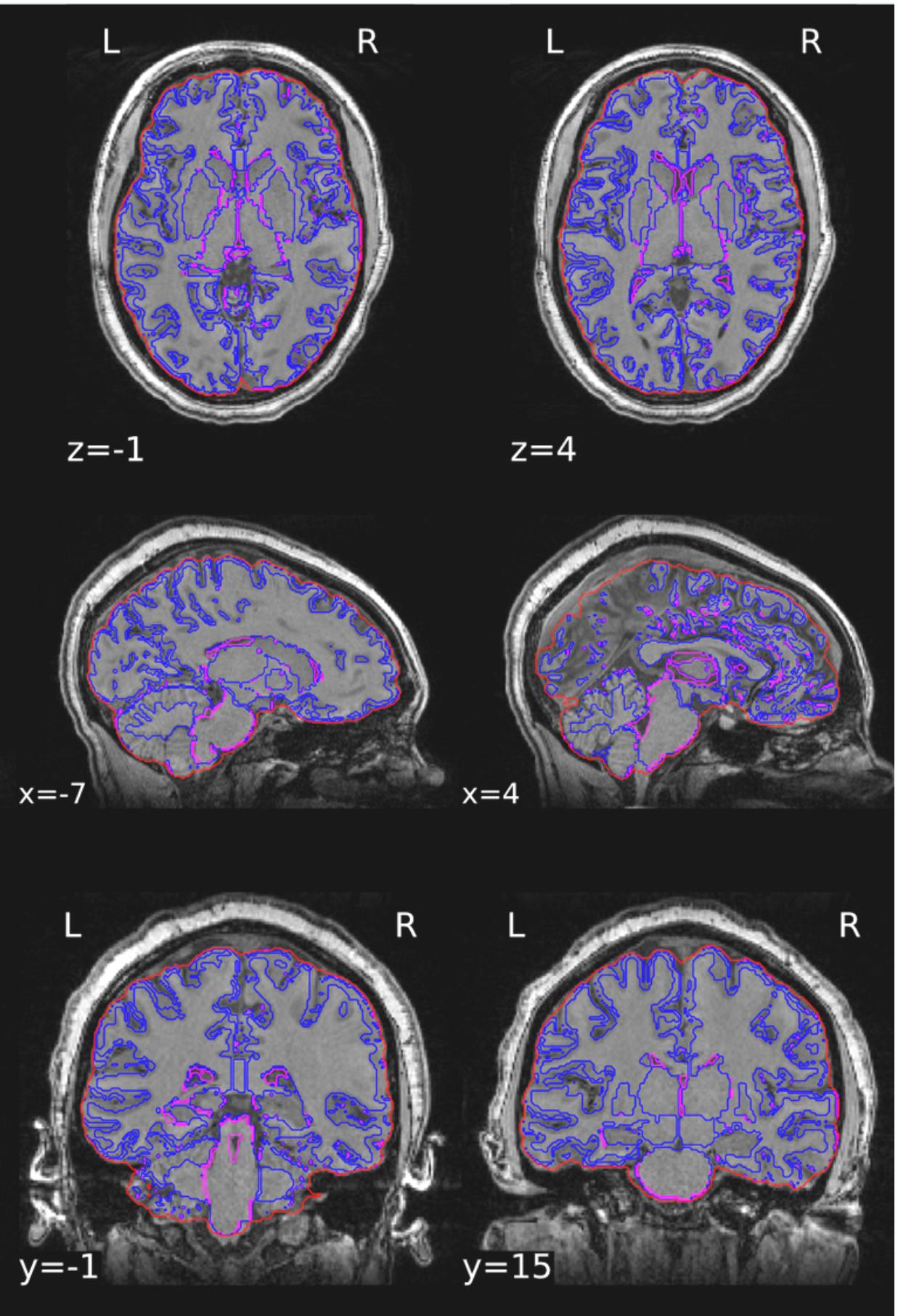
- Uses *antsBrainExtraction.sh* from ANTs
- Atlas-based - registers atlas whole-head image to T1w and applies atlas's brain mask to create subject's brain mask
- Sometimes referred to as "skull-stripping"



Tissue Segmentation

Divides image into tissue types (white matter, gray matter, CSF)

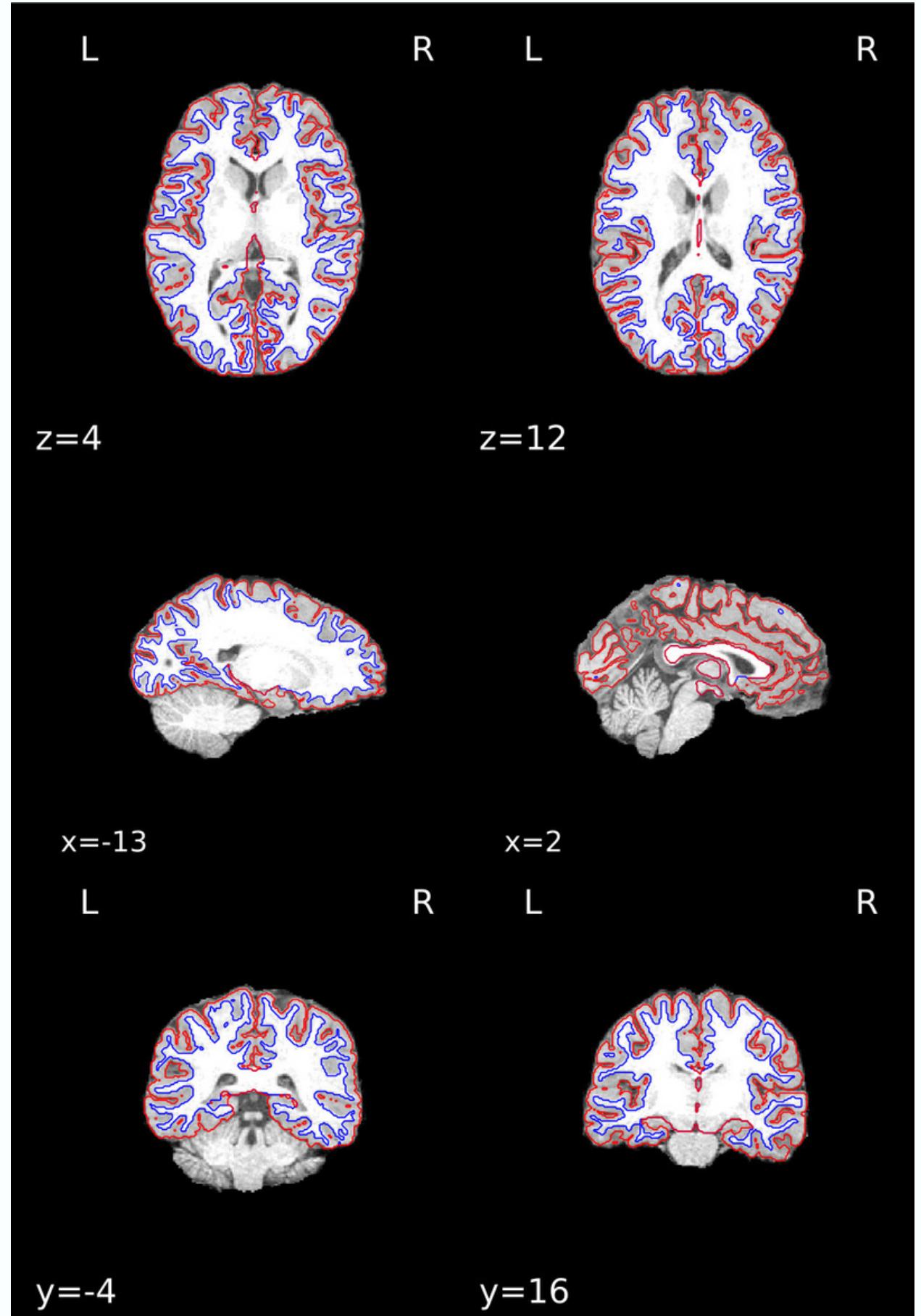
- Uses *fast* from FSL
- Model classifies tissue based on the voxel's intensity and the intensity of neighboring voxels
- Reciprocally incorporates/assesses magnetic field inhomogeneity



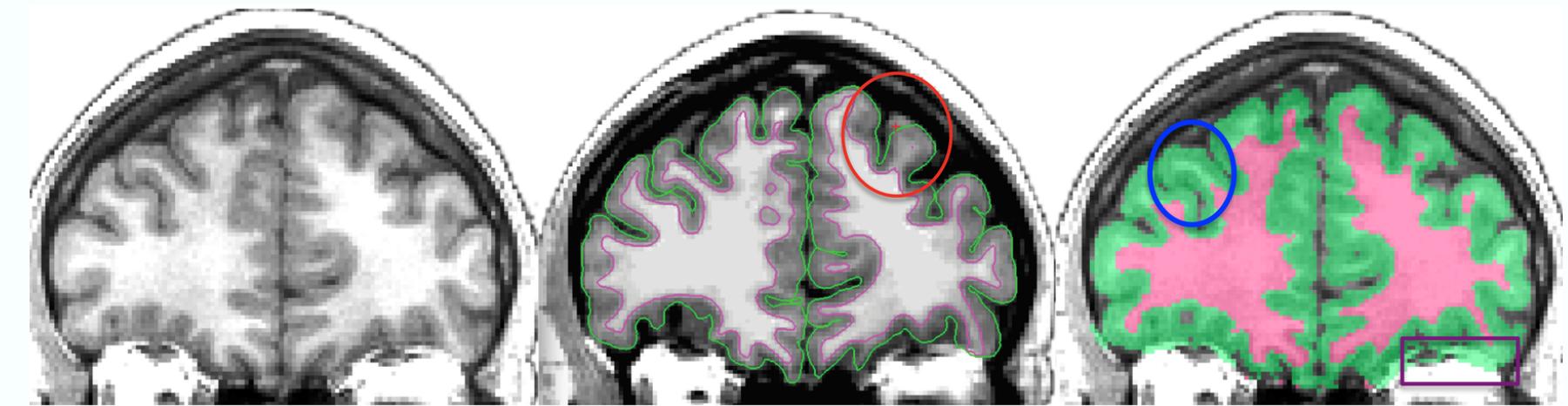
Surface Preprocessing

Optional assessment of boundary between white and gray matter (wm surface) and the cortical surface (pial surface)

- Uses *recon-all* from FreeSurfer
- Also calculates *.midthickness surfaces
- Saves surfaces in GIFTI format
- Uses T2w image if available



Refine Brain Mask



Klein et al, 2017

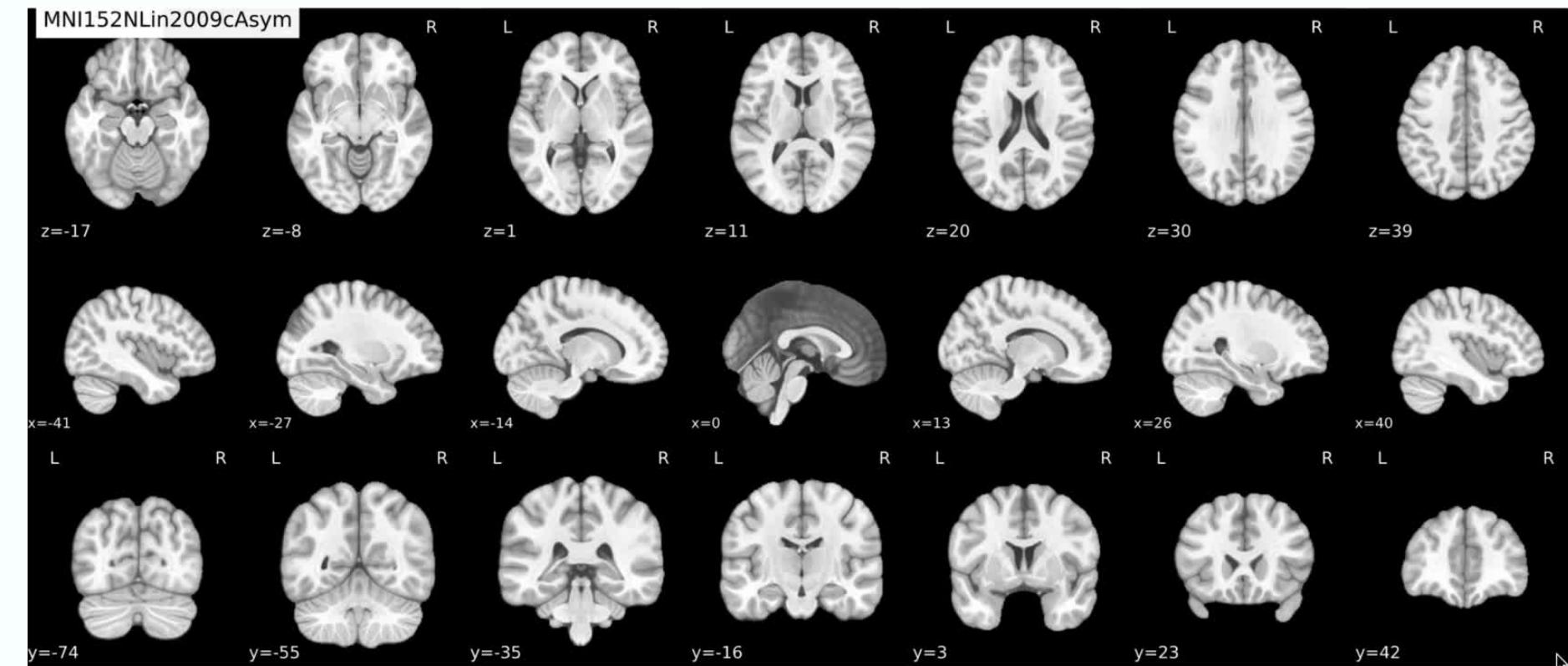
**Cleans up the brain mask calculated by
brain extraction based on the surface
segmentation**

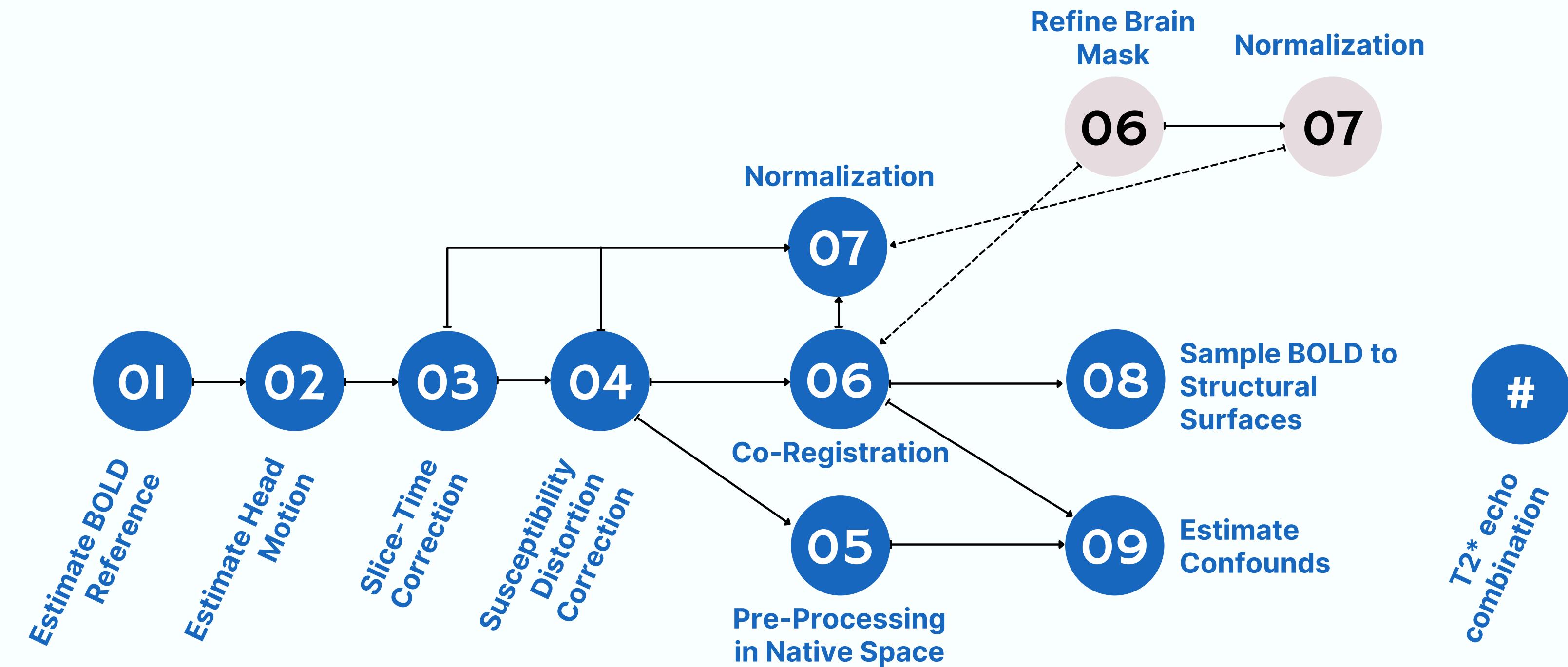
- Uses a custom adaptation of Mindboggle's hybrid segmentation algorithm
- Unifies ANTs and FreeSurfer segmentations for the most accurate results

Normalization

Fits subject's anatomical image to a standard template so you can compare between individuals

- Uses *antsRegistration* from ANTs
- Non-linear resampling
- Can chose from standard or nonstandard templates ("spaces")
- Keeps original resolution (unless specified)

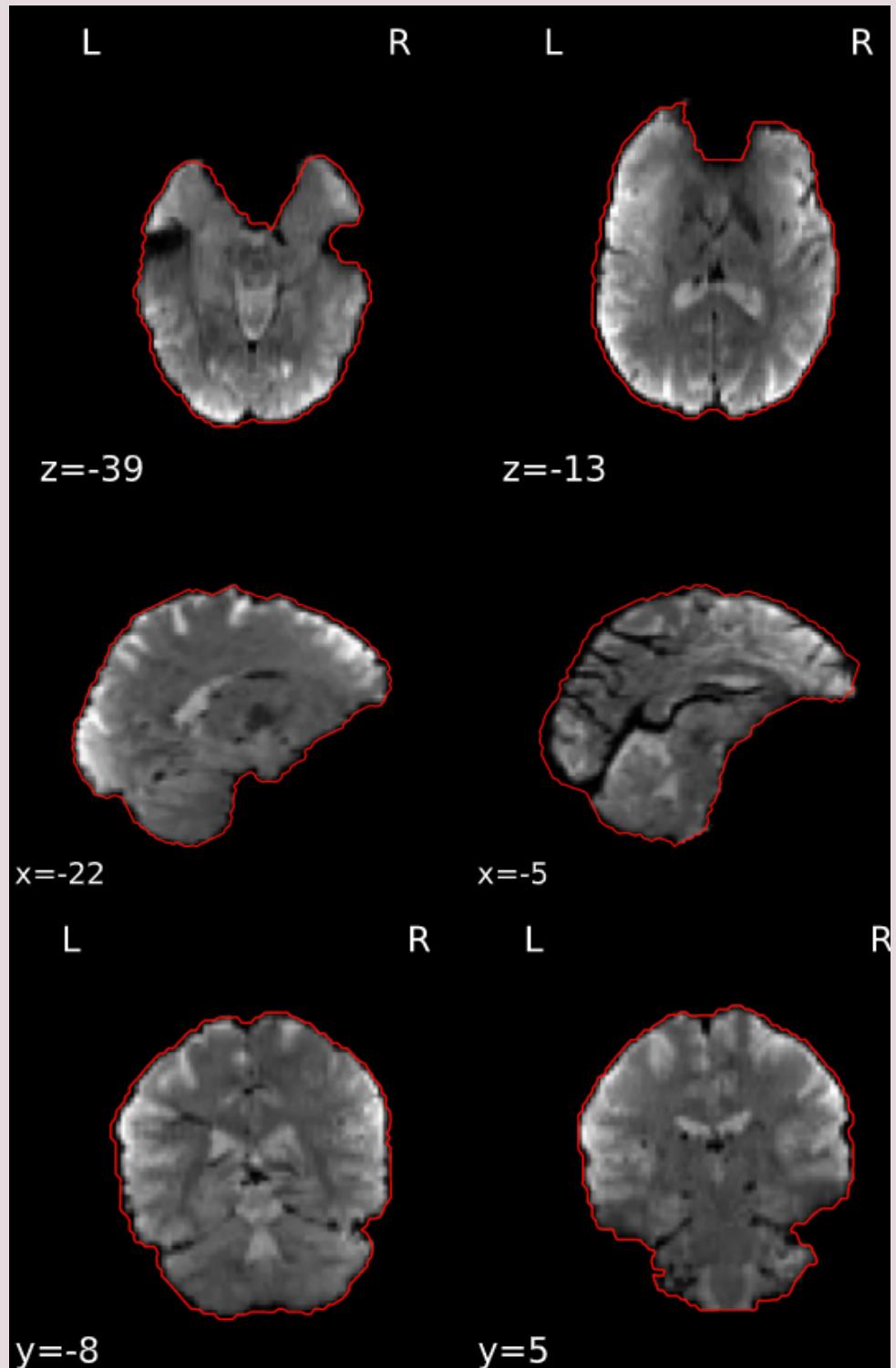




Estimate BOLD Reference

Creates a reference image from each functional run by averaging dummy scans or a motion-corrected subset of BOLD volumes

- Also creates BOLD run brain mask
- Uses sbref (single-band reference) if available

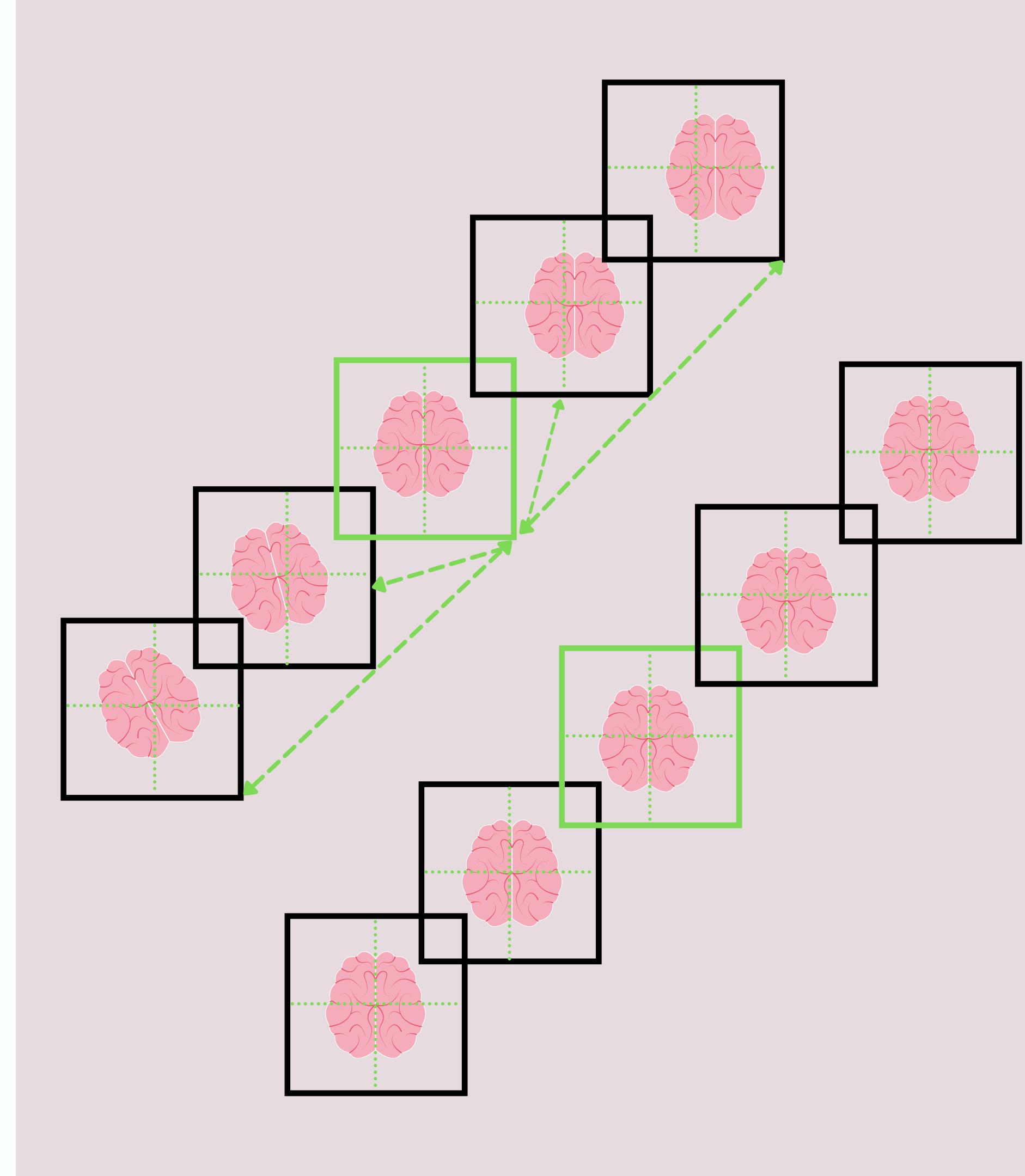


fMRIPrep Sample Report

Estimate Head Motion

Writes a rigid-body transform - steps needed to bring brain back into alignment with the reference image - for each volume

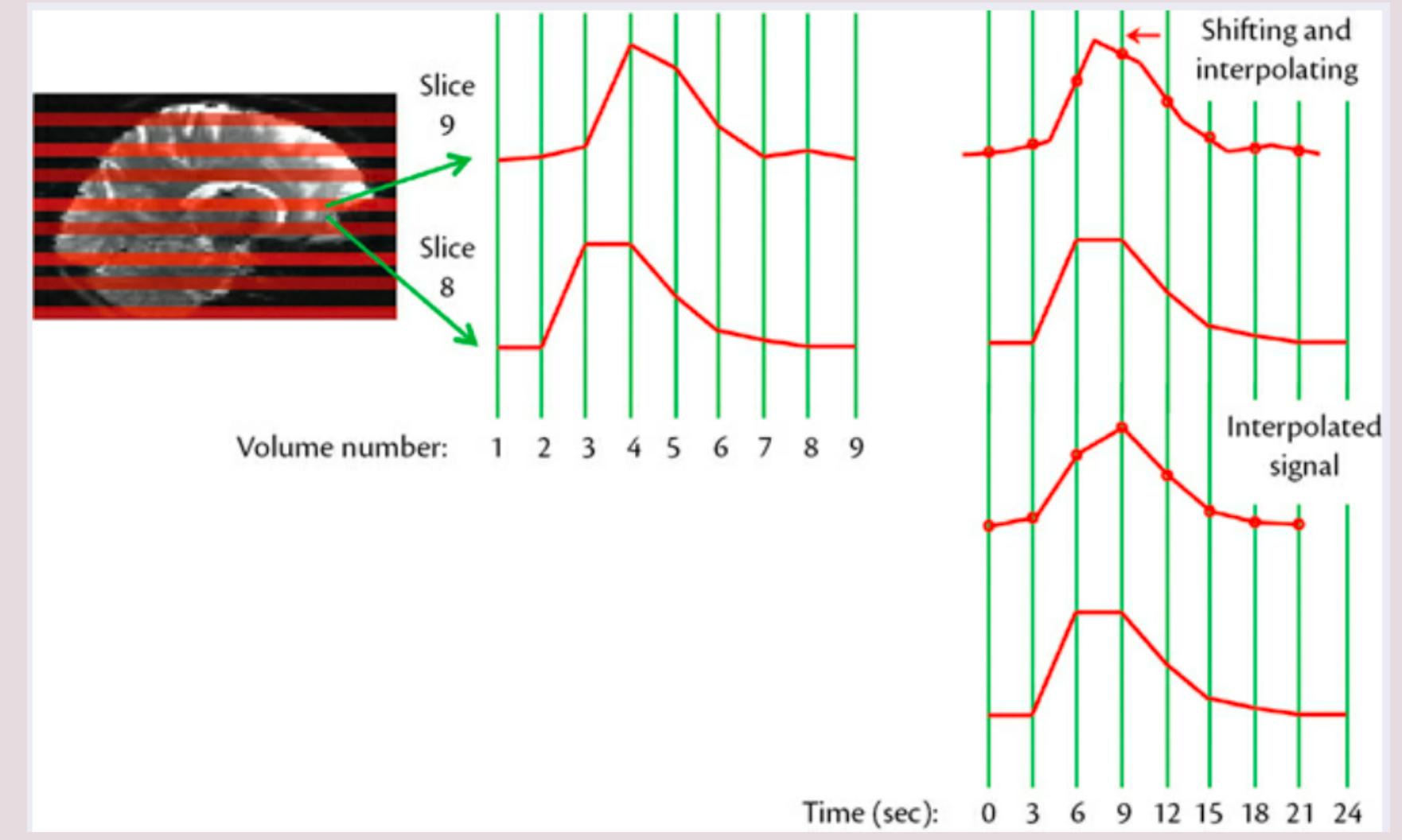
- Uses *mcflirt* from FSL
- Calculated from 6 motion parameters (3 rotations & 3 translations) per time-step



Slice Time Correction

Realigning 2D image slices to the middle TR (i.e. middle of that volume's acquisition)

- Uses *3dTShift* from AFNI
- Can be skipped

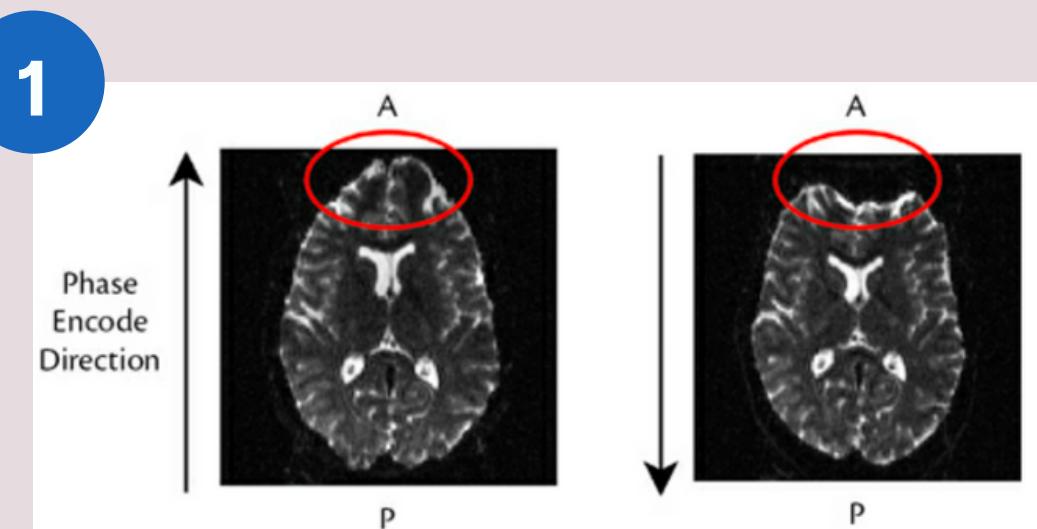


Jenkinson & Chappell, 2018

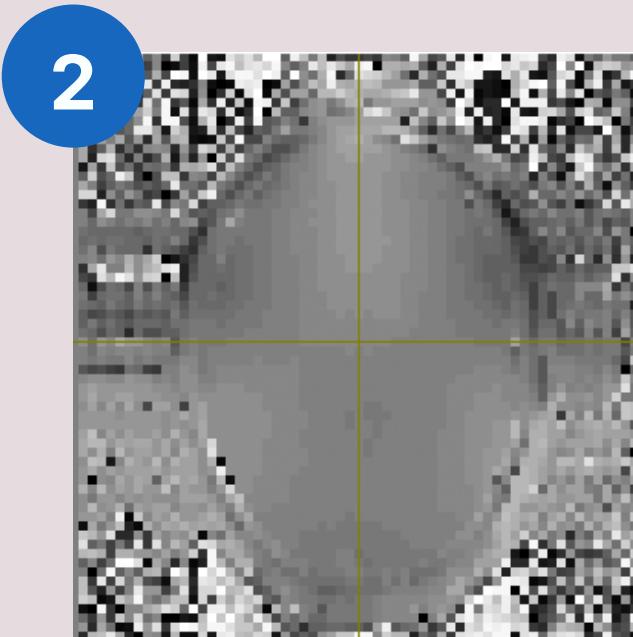
Susceptibility Distortion Correction

Correcting signal intensity noise caused by inhomogeneities in the magnetic field

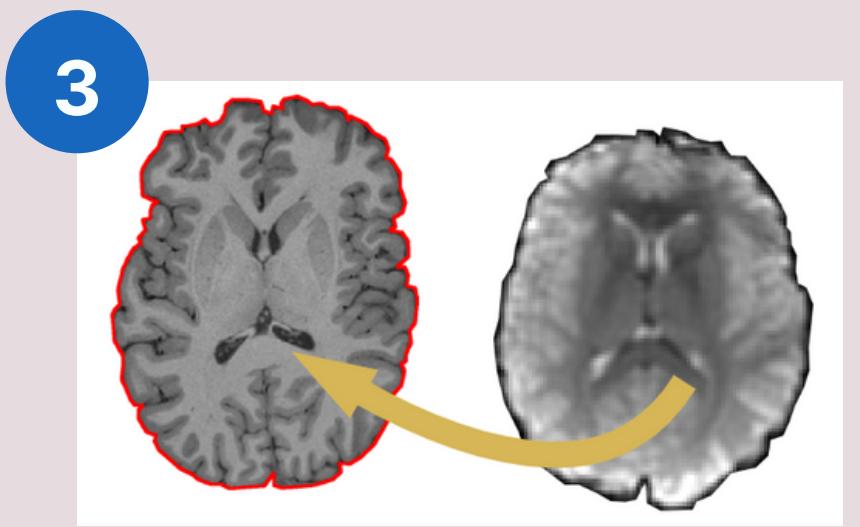
- Calculates the fieldmap by:
 1. PE-Polar: Registering BOLD images acquired with opposite phase encoding directions to a reference and calculating voxel displacement
 2. Direct: comparing echos in a double-echo sequence to reveal inhomogeneities
 3. Fieldmap-less: Registering BOLD images to T1w to assess voxel displacement



Jenkinson & Chappell, 2018

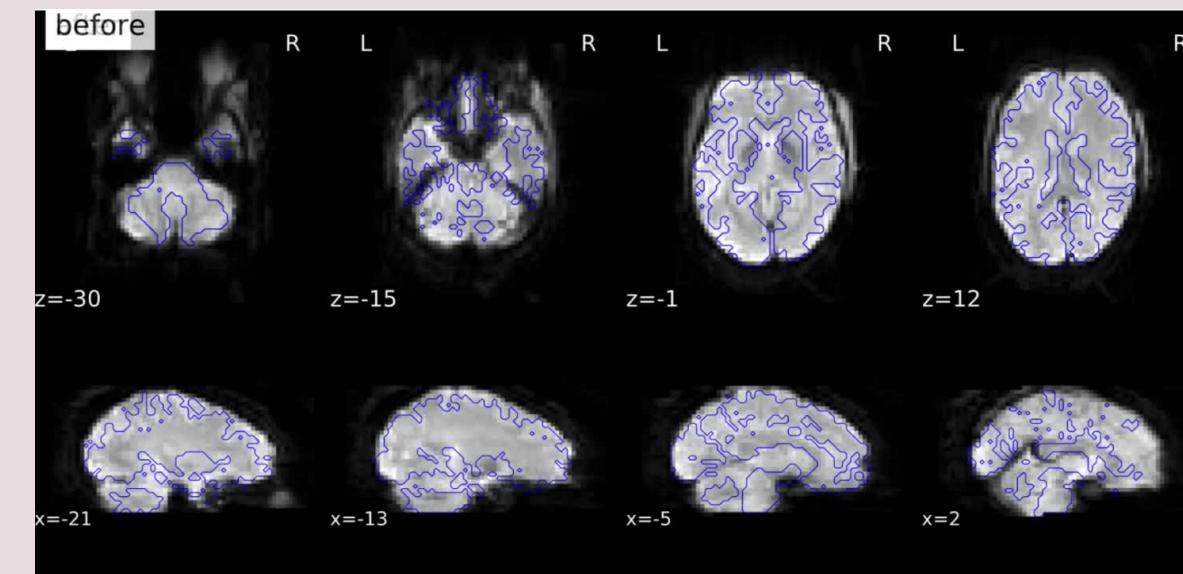


2



3

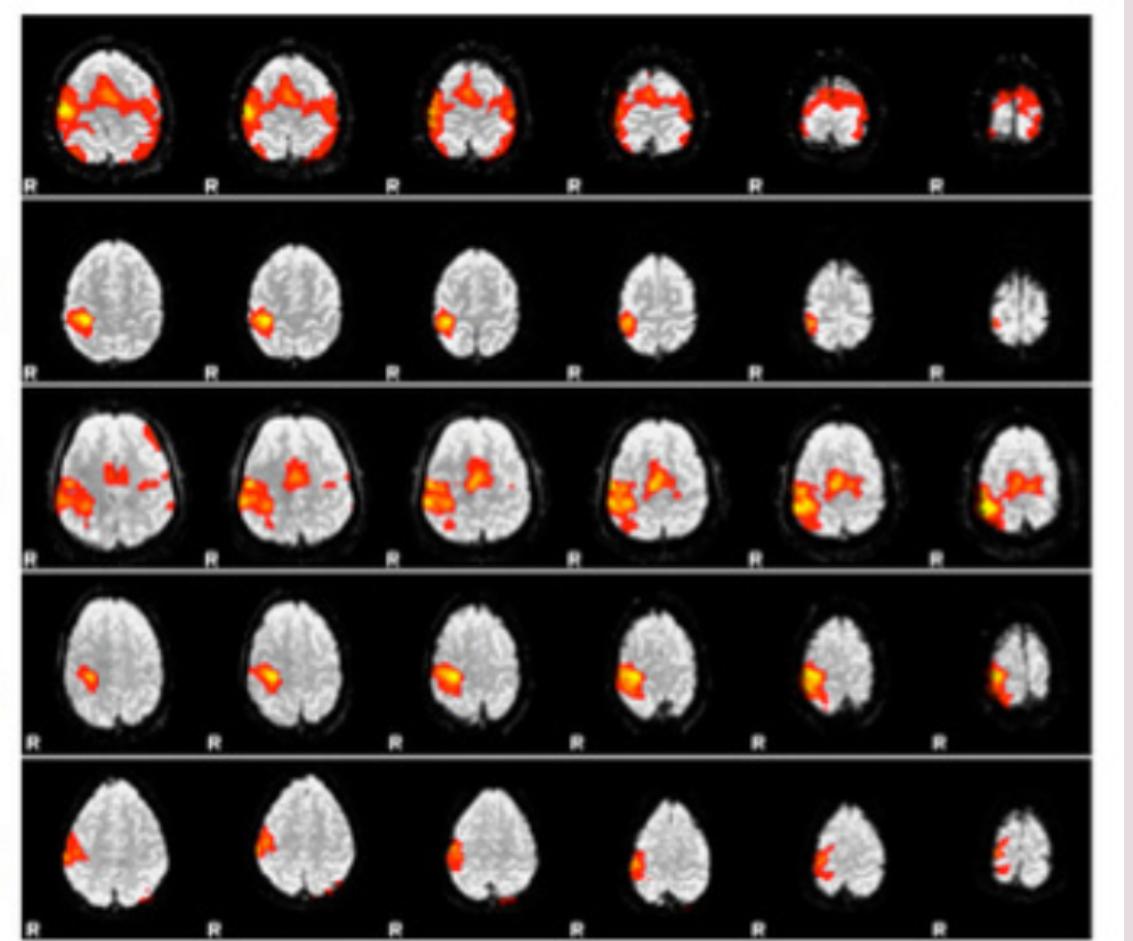
Adapted from Esteban et al, 2019



Pre-process BOLD in Native Space

Applying transforms calculated by
**motion correction, slice-timing
correction, and signal distortion
correction**

Subject 1
Subject 2
Subject 3
Subject 4
Subject 5

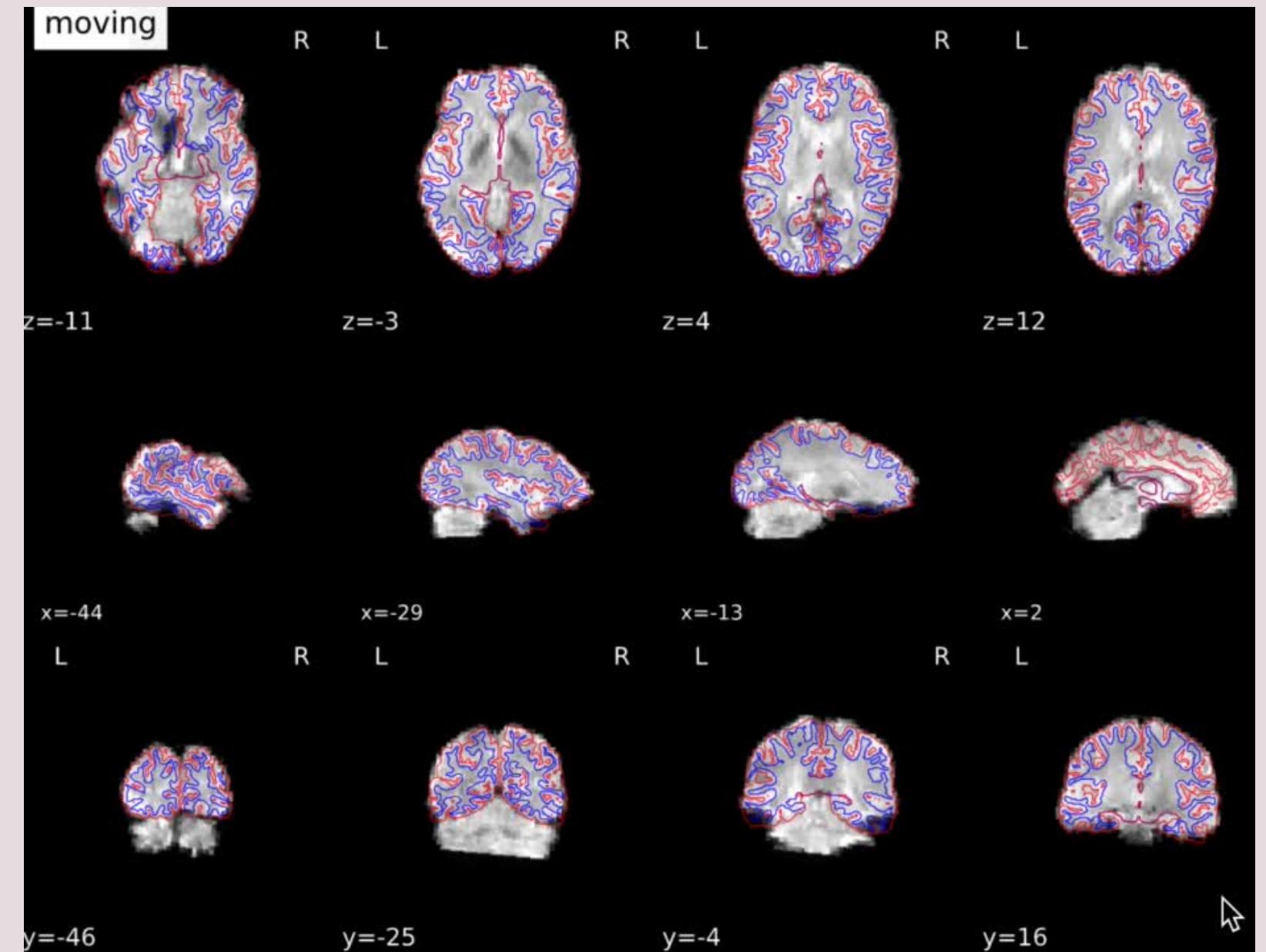


Adapted from [MRI Questions.com](https://www.mriquestions.com)

Co- Registration

Aligns the BOLD reference image to the T1w brain mask by maximizing it's intensity gradient across the white-matter boundary

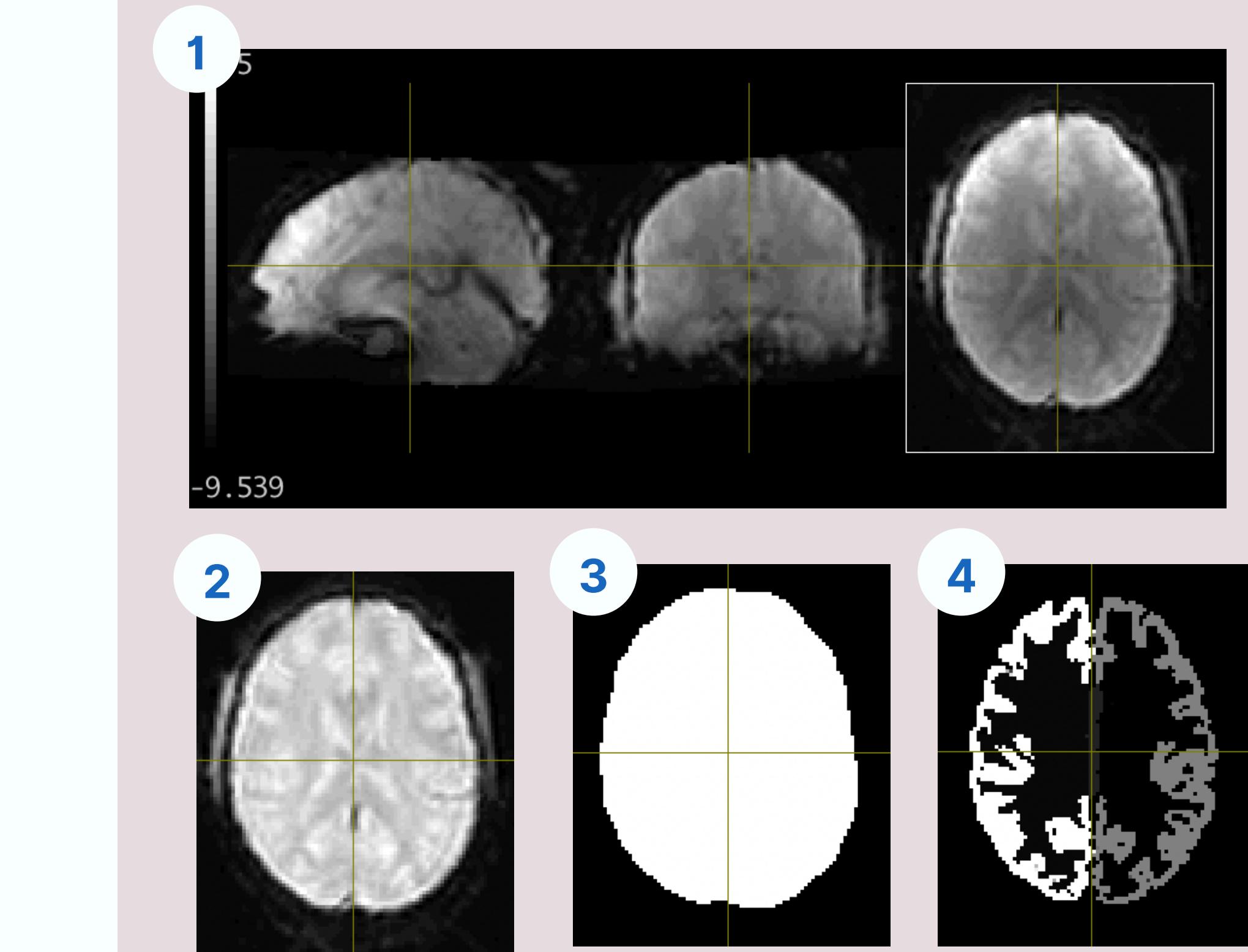
- Uses *bbregister* from FreeSurfer or *FLIRT* from FSL



Normalization

Concatenates and applies all the transforms to map BOLD and T1w onto standard space

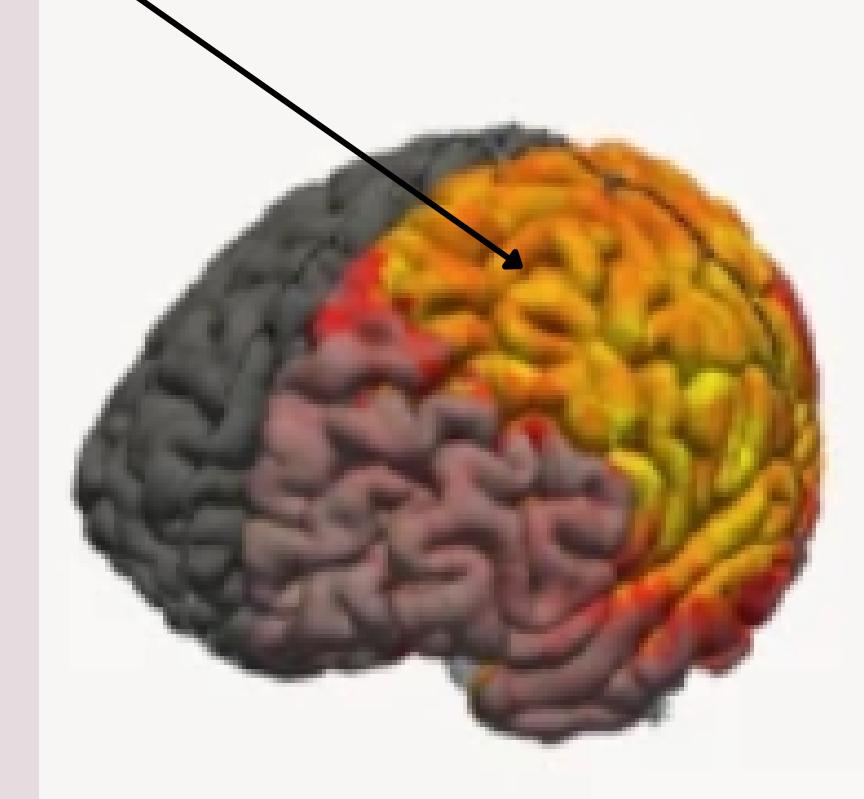
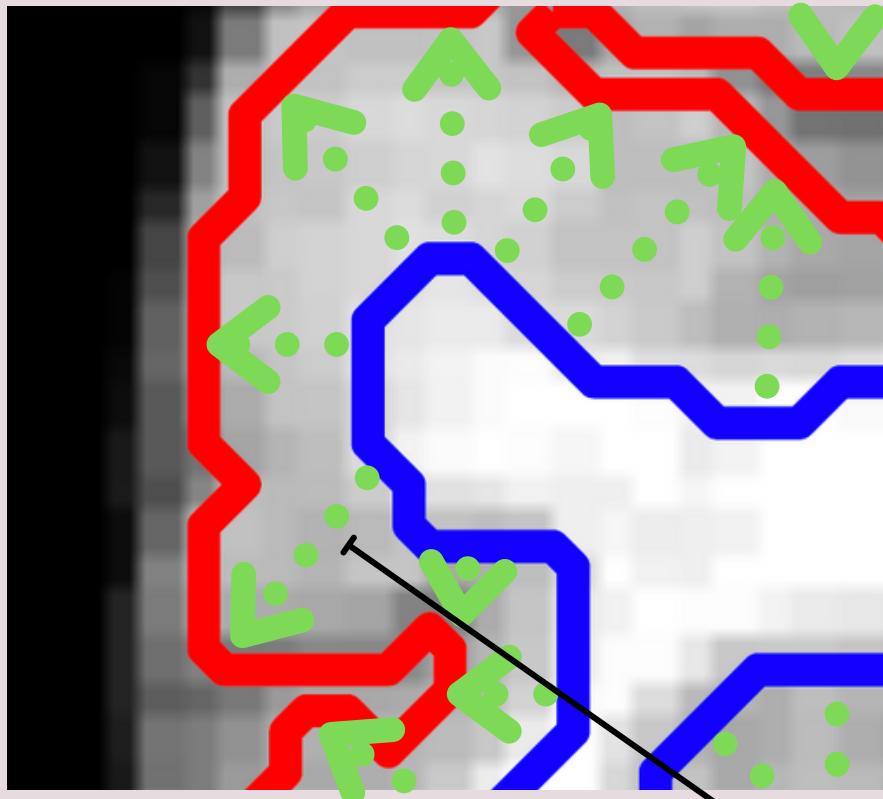
- Simultaneously applies the transforms calculated by Head Motion Estimation, Susceptibility Distortion Correction, Co-Registration, and T1w Normalization
- Outputs normalized:
 - 1.fMRI (BOLD) series
 - 2.BOLD reference
 - 3.T1w brain mask
 - 4.FreeSurfer surfaces



Sample BOLD to Structural Surfaces

BOLD timeseries is aligned to the cortical surfaces by averaging across the cortical ribbon

- Uses *mri_vol2surf* from FreeSurfer
- Calculated for native space and fsaverage (FreeSurfer's standard space)

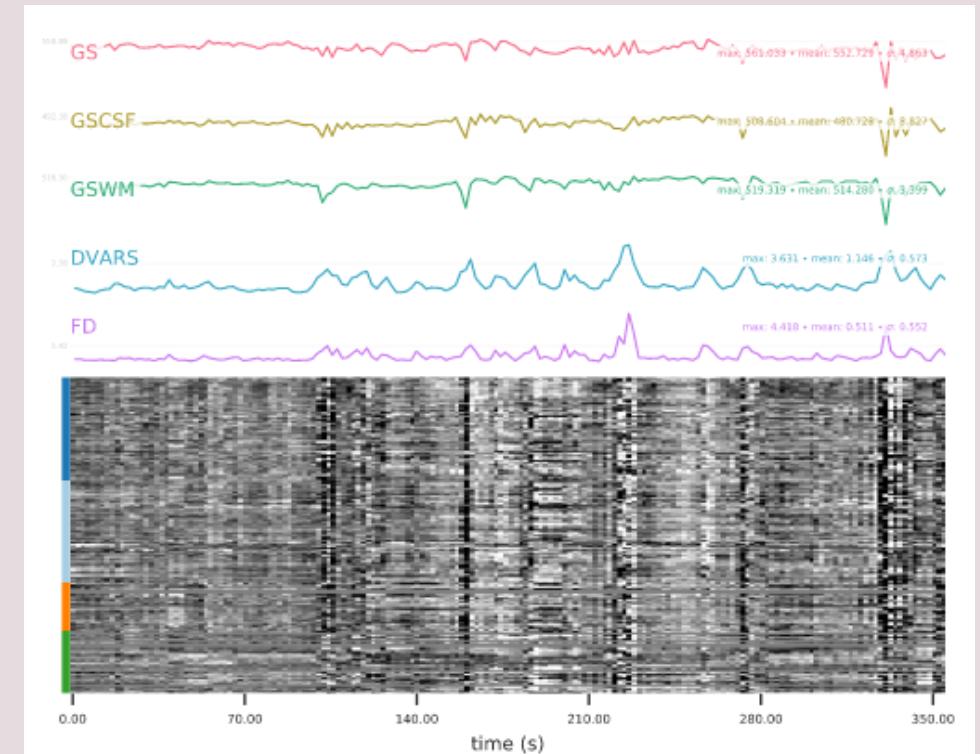
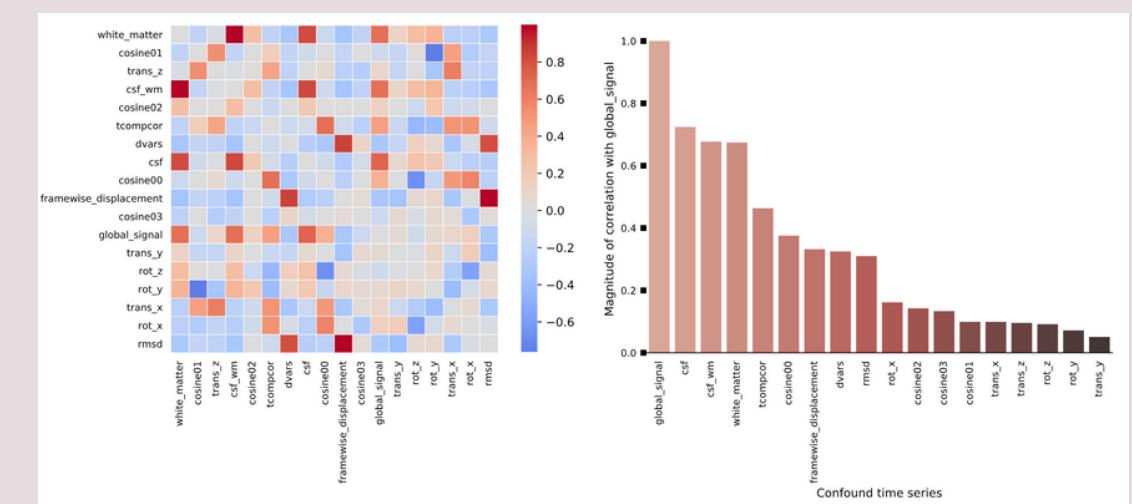
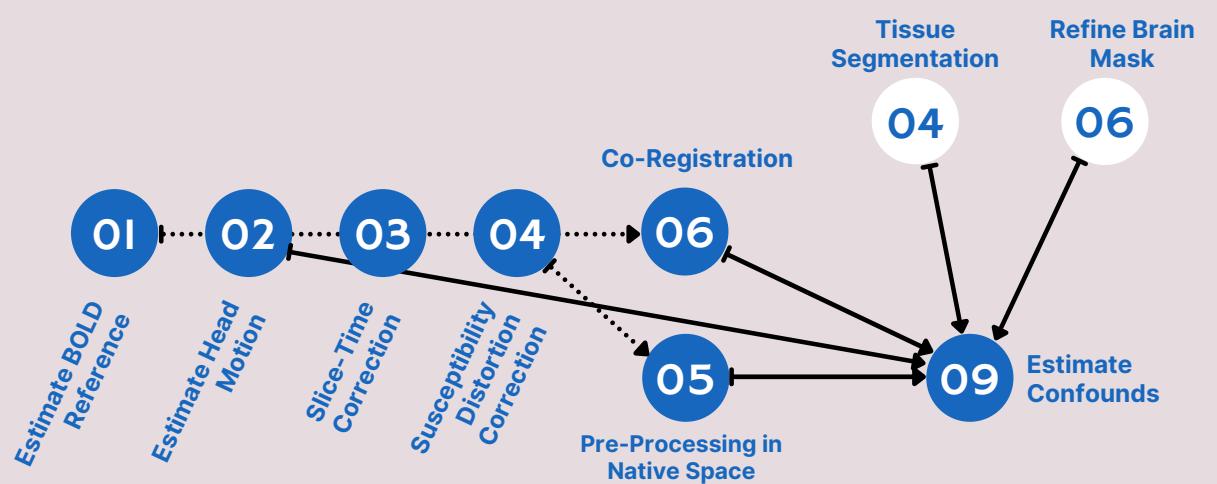


Adapted from Esteban et al, 2019

Estimate Confounds

Identifies many sources of potential noise in the signal that you can choose if/how to resolve:

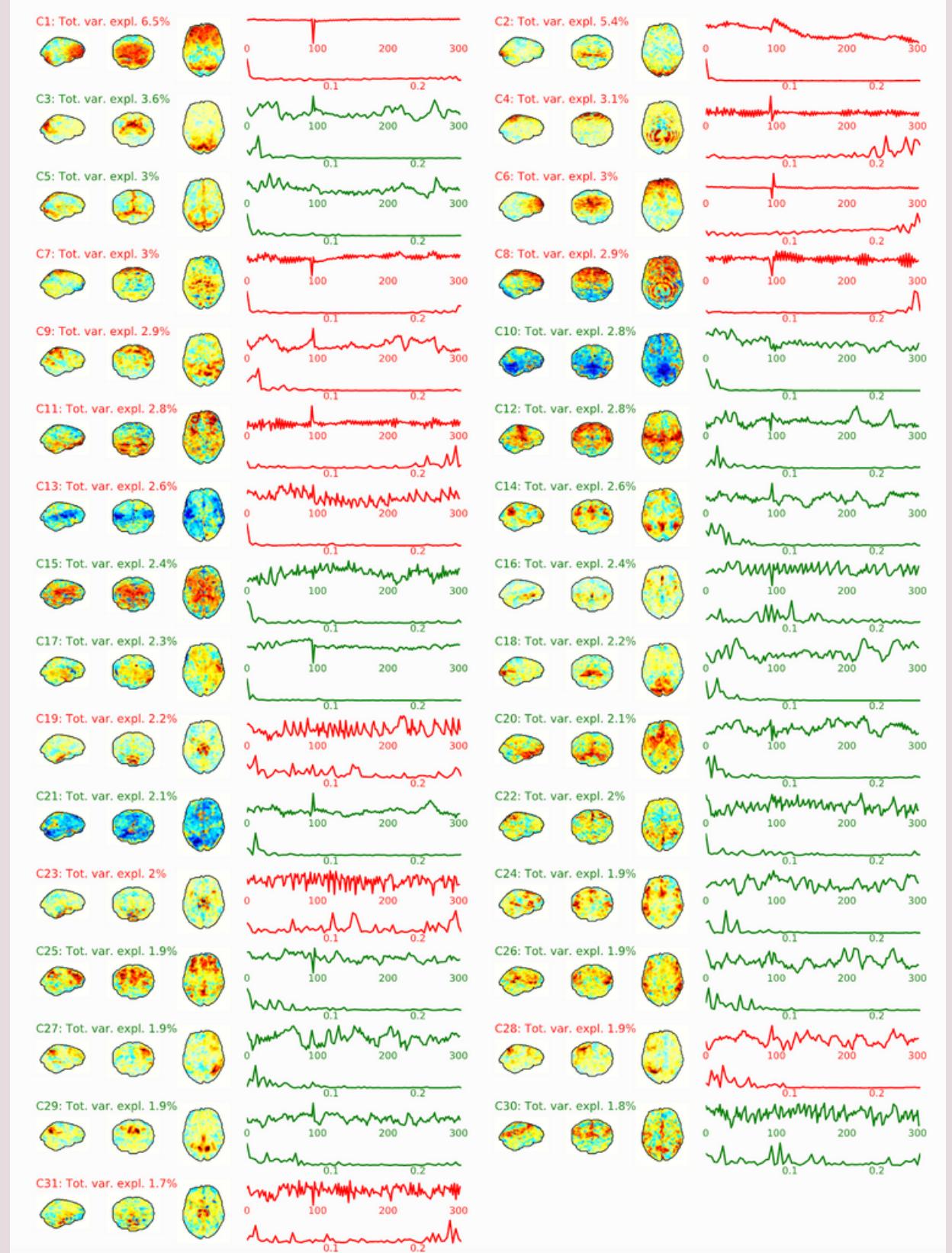
- Mean signals (global and for each tissue type)
 - Frame-wise Displacement
 - 6 motion parameters
 - Spike regressors
 - Temporal and Anatomical CompCore
 - DVARS
 - ICA-AROMA



ICA-AROMA

**Sub-workflow of Confound Regression
that uses independent component
analysis (ICA) to identify and remove
signal caused by head motion**

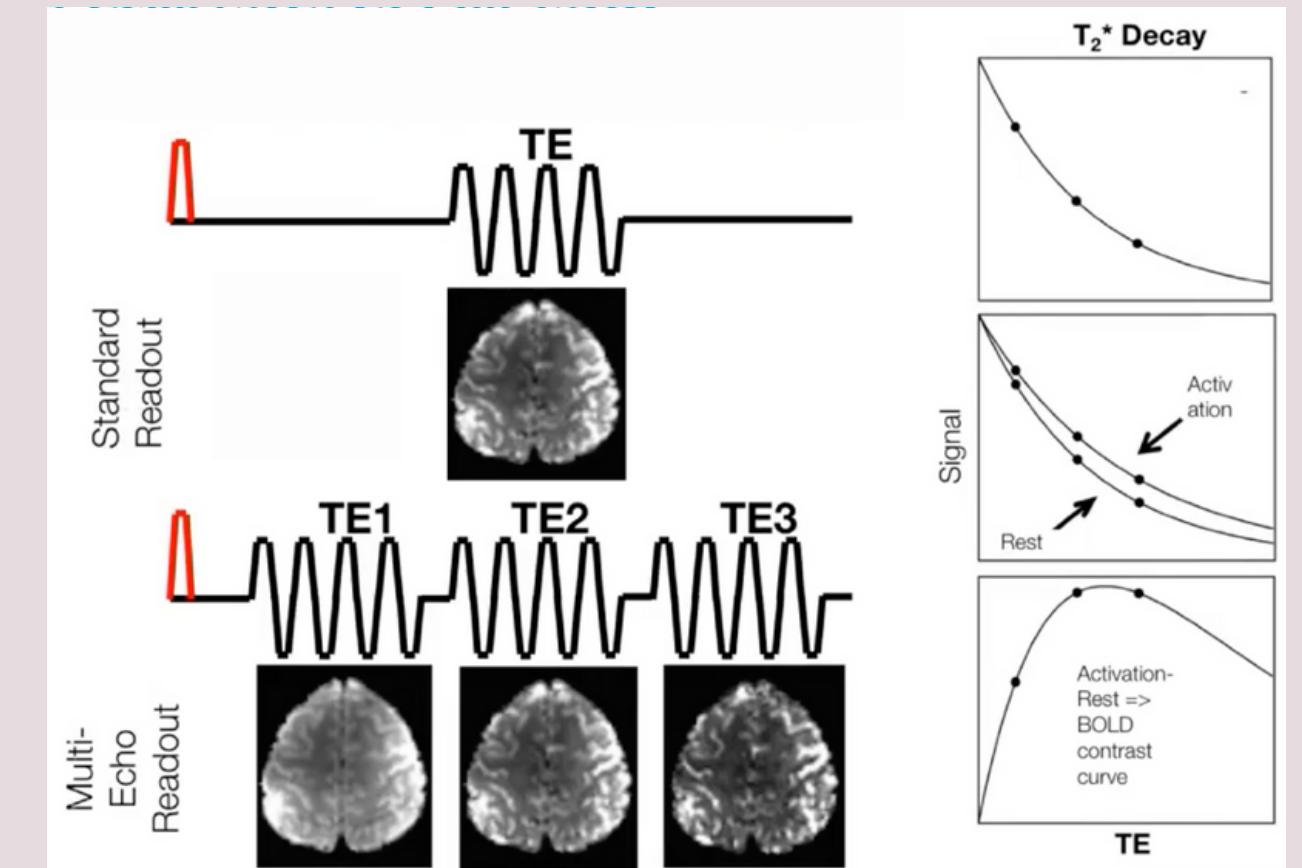
- Applies non-aggressive denoising to normalized BOLD data
- Calculated for both native space and normalized BOLD
- Provides confounds to run aggressive denoising if desired



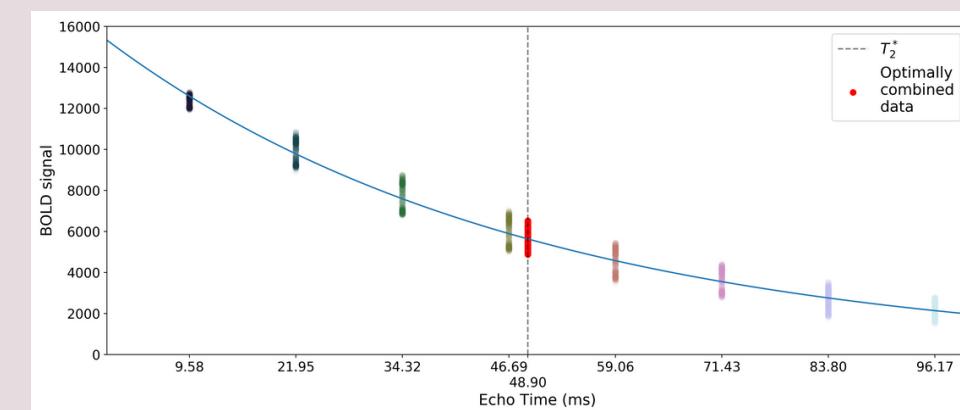
T₂*-driven Echo Combination

Uses multi-echo data to create a T₂* map then optimally combines echos into a single BOLD time series

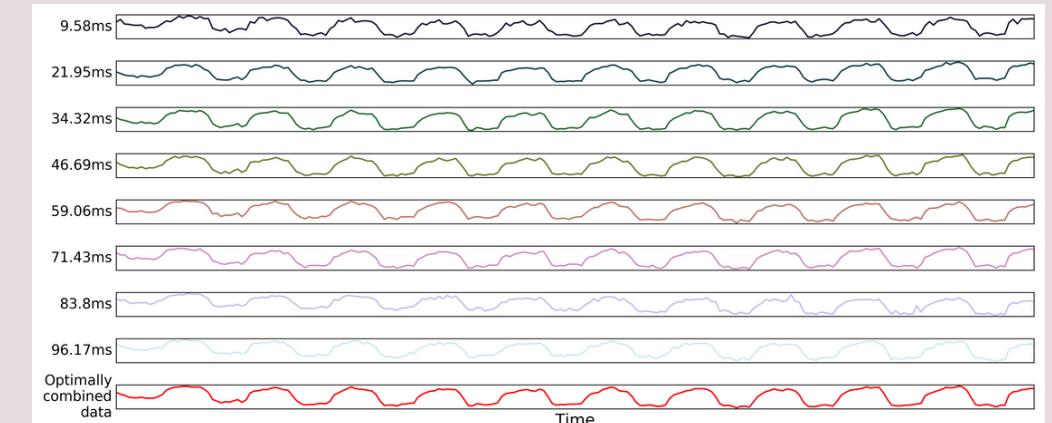
- Integration of part of tedana
- Estimates T₂* for each voxel to create a map
- Uses a weighted average to compute BOLD at each voxel's optimum TE



Adapted from Kundu, P, 2017



tedana developers, Revision 4d5c6456



Thanks!

Tinashe Tapera

Sydney Covitz

Matt Cieslak

Azeez Adebimpe

Adam Pines

Dan Wolf

Ted Satterthwiate

and everyone for listening!



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