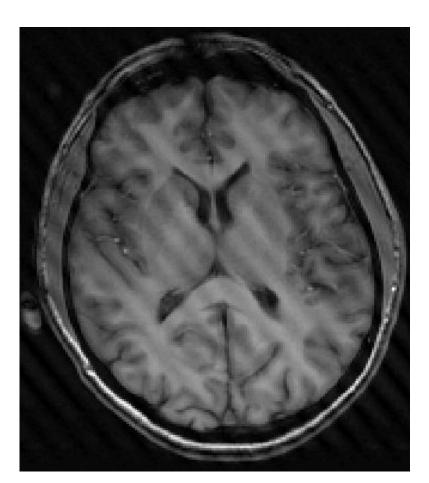


# FMRI single subject analysis

- Overview
- fMRI experiment basics
- Preprocessing
  - motion correction
  - slice timing
  - temporal filtering
  - smoothing



~ MM Mahaman MM



# Generic study blueprint



1. Data acquisition

2. Data preprocessing

3. Single-subject analysis

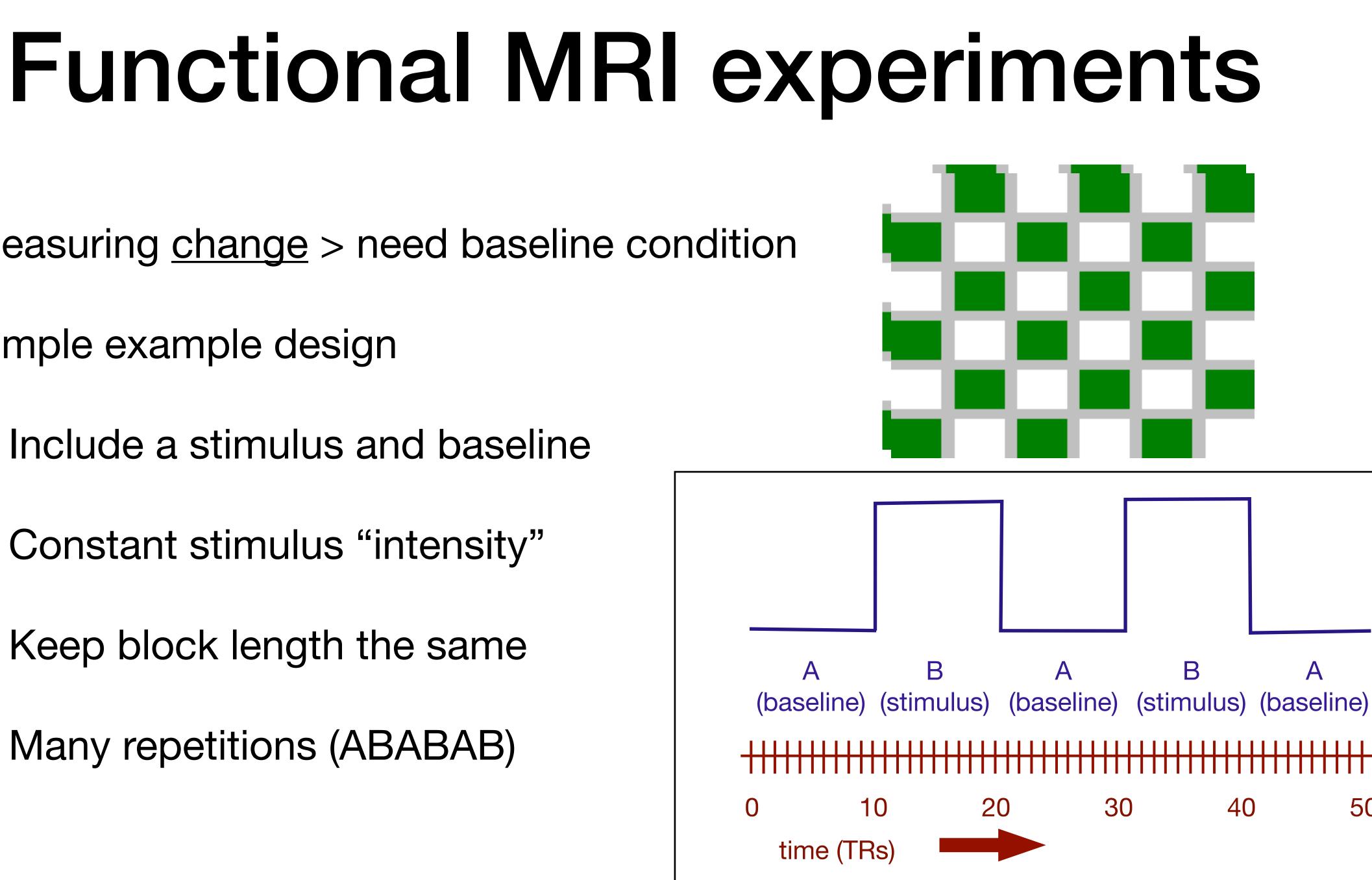
4. Group-level analysis

5. Statistical inference





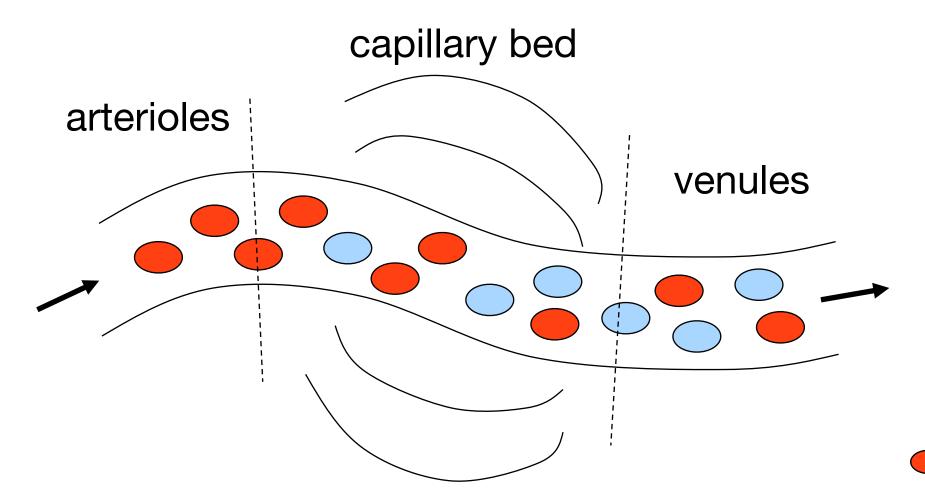
- Measuring <u>change</u> > need baseline condition
- Simple example design
  - Include a stimulus and baseline
  - Constant stimulus "intensity"
  - Keep block length the same
  - Many repetitions (ABABAB)







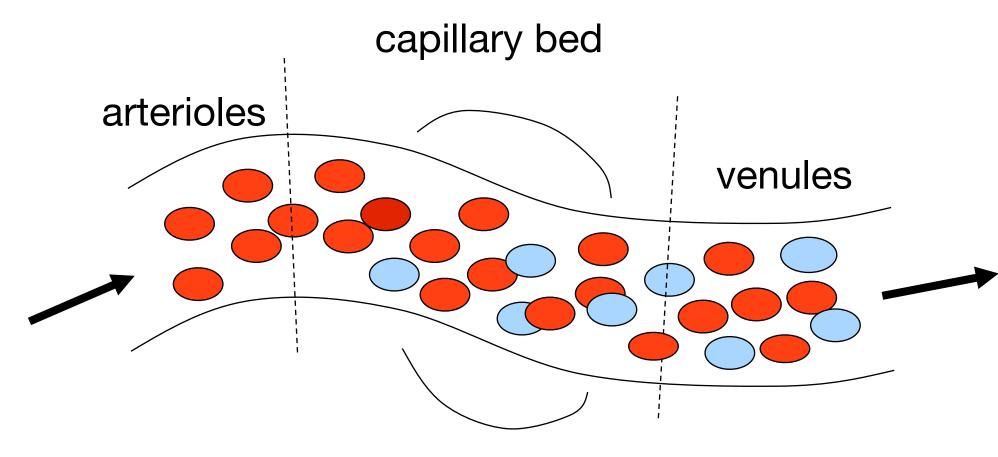
## The Hemodynamic Response



#### **Basal State**



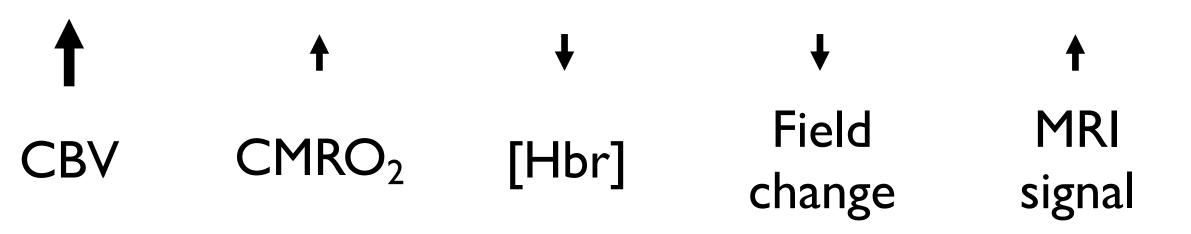
CBF



- = HbO2
- = Hbr

 $\bigcirc$ 

#### **Activated State**

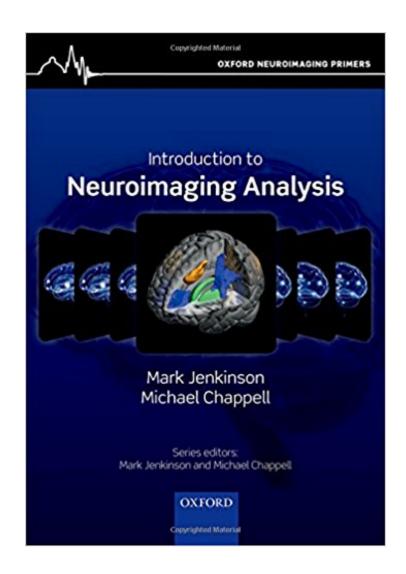


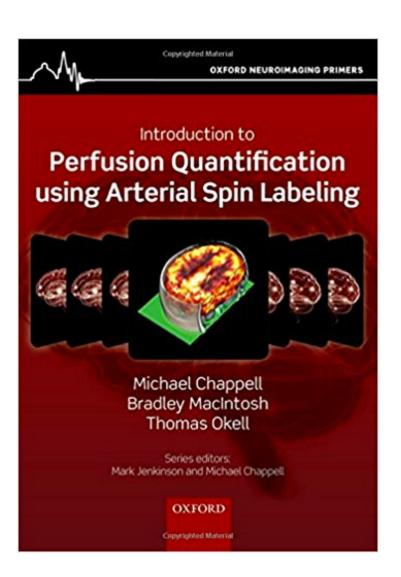
BOLD MRI =  $T_2^*$ -weighted



## The free online appendix

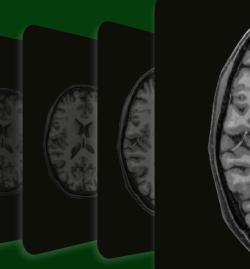
- Part of a series of Oxford Neuroimaging Primers
- <u>https://www.fmrib.ox.ac.uk/primers/appendices/</u> mri physics.pdf

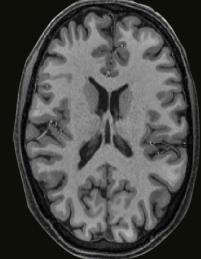


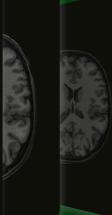




#### Short introduction to **MRI** Physics for Neuroimaging







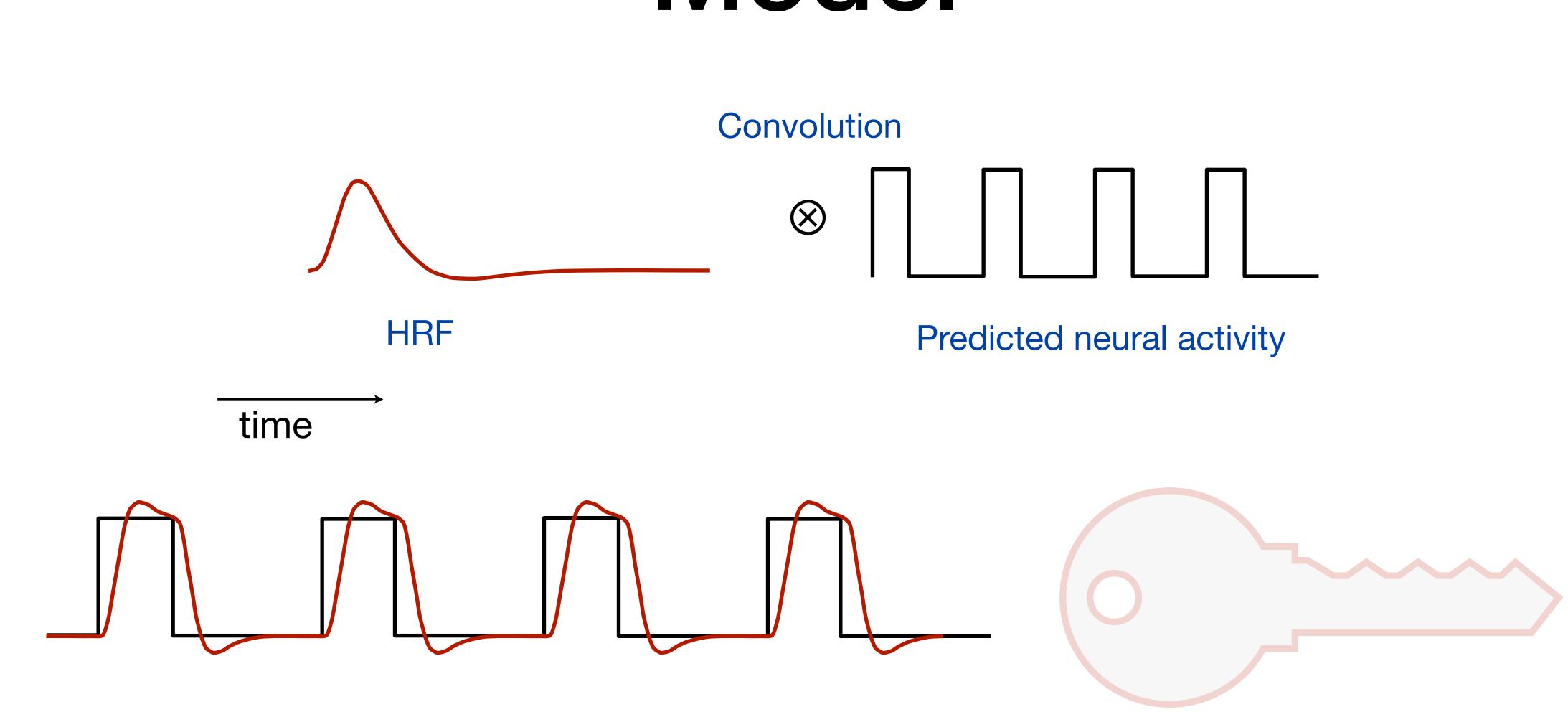
Michael Chappel **Thomas Okell** Mark Jenkinson

Series editors: Mark Jenkinson and Michael Chapp

> PRIMER **APPENDIX**

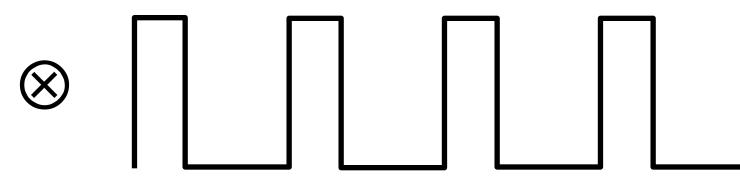






Predicted response

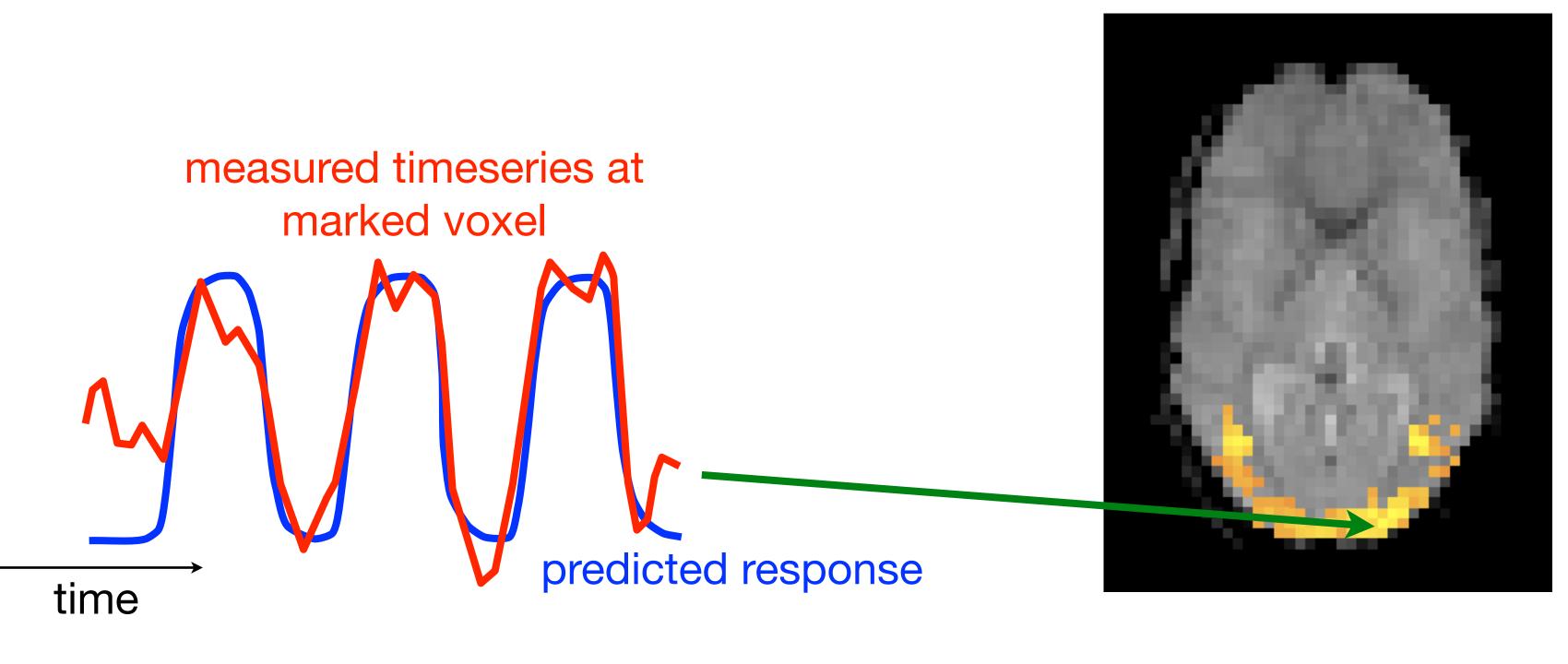
### Model





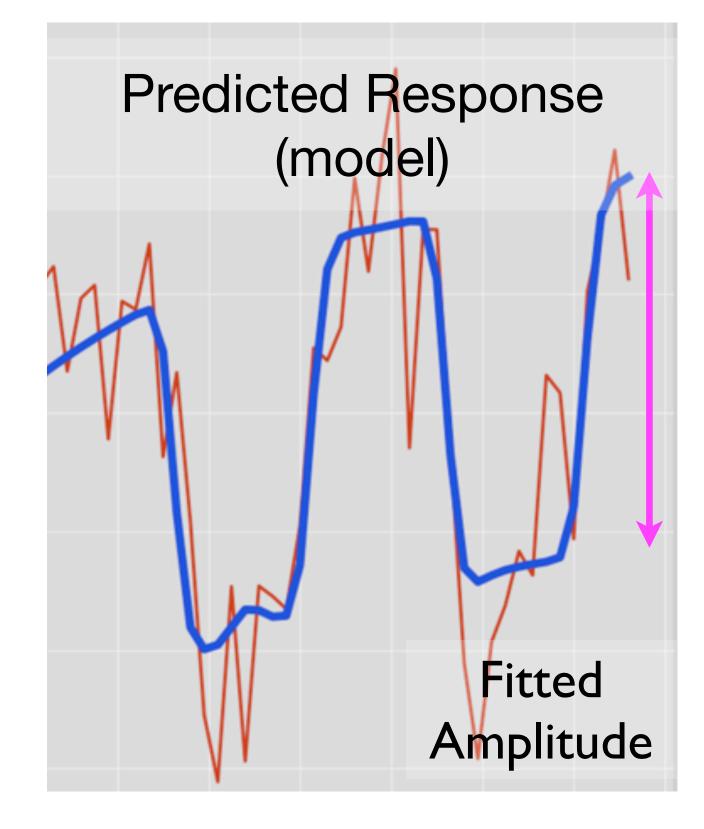
### Fit model to data

Look for voxels that have a BOLD timeseries similar to the model



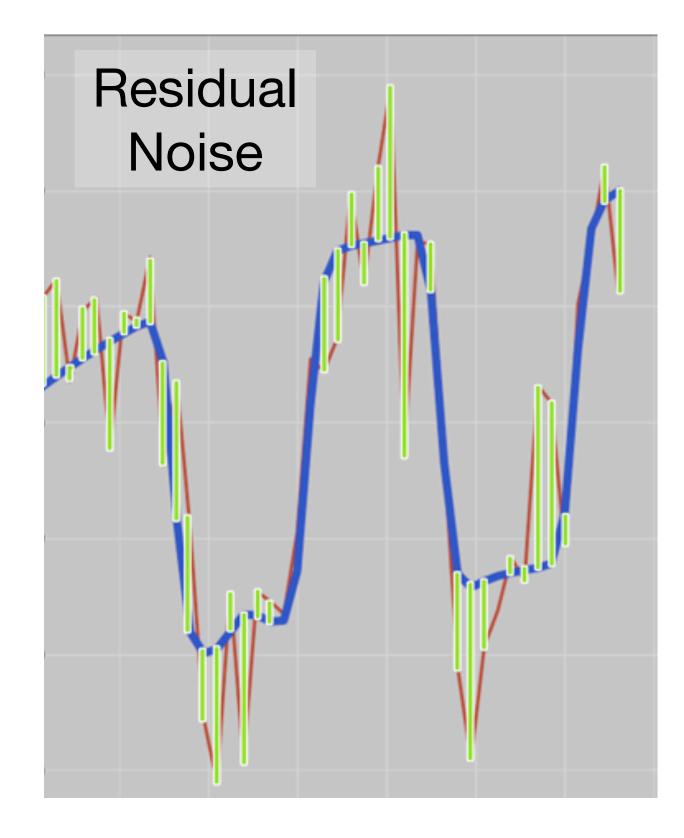


• Fit model to each voxel separately



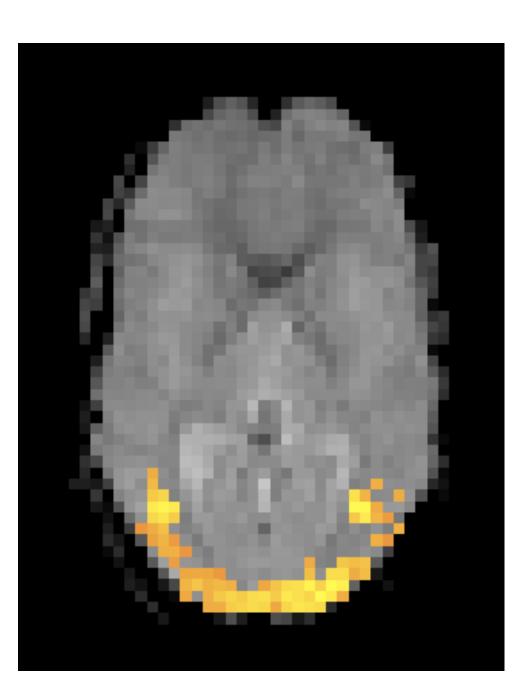


- Fit model to each voxel separately
- Measure residual noise variance





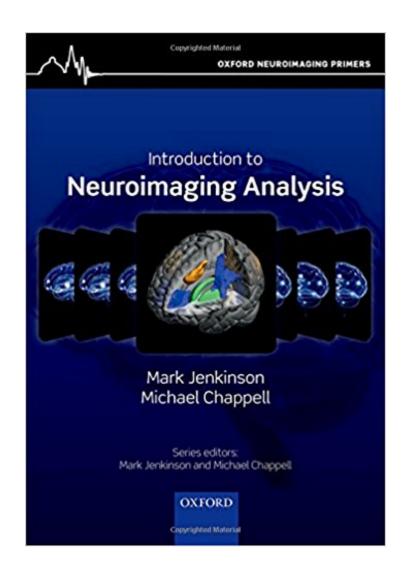
- Fit model to each voxel separately
- Measure residual noise variance
- Voxel t-statistic = model fit / noise amplitude
- Threshold t-statics and display map

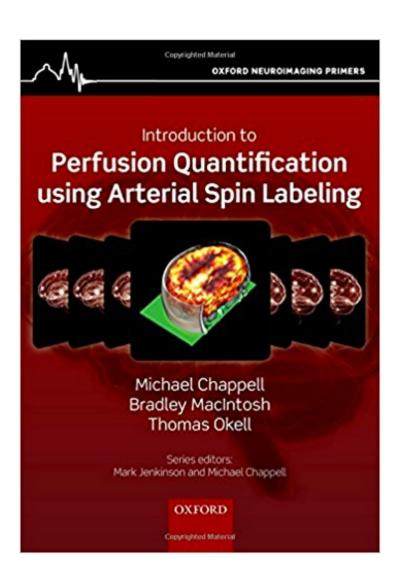


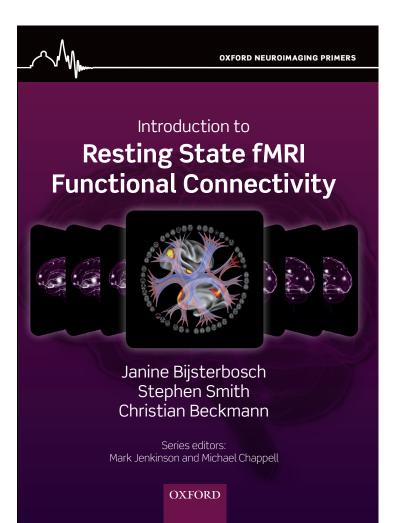


## The free online appendix

- Part of a series of Oxford Neuroimaging Primers
- https://www.fmrib.ox.ac.uk/primers/appendices/ lacksquare<u>glm.pdf</u>

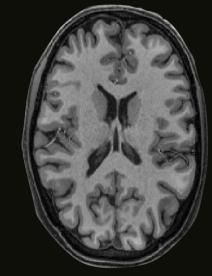






#### Short introduction to the **General Linear Model** for Neuroimaging

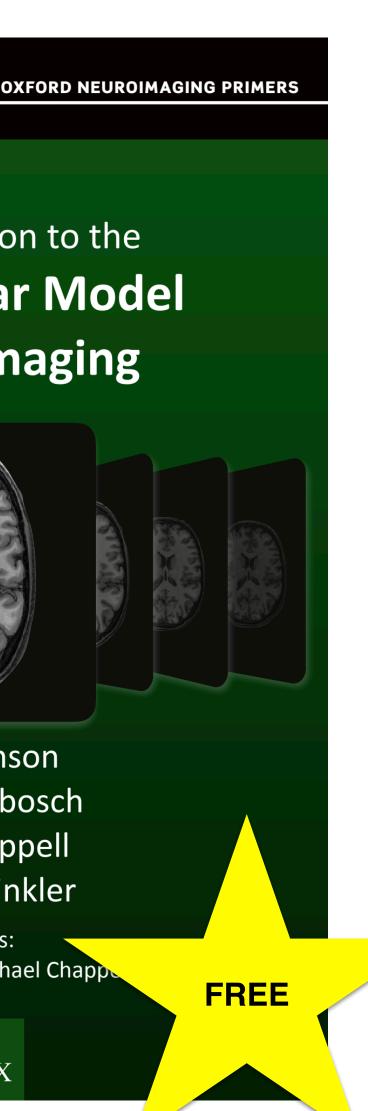




Mark Jenkinson Janine Bijsterbosch Michael Chappell Anderson Winkler

Series editors: Mark Jenkinson and Michael Chappe

> PRIMER APPENDIX

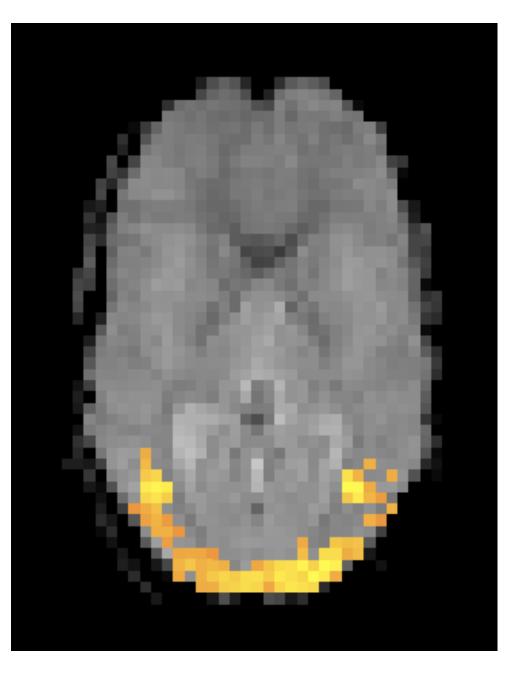




- Fit model to each voxel separately
- Measure residual noise variance
- Voxel t-statistic = model fit / noise amplitude
- Threshold t-statics and display map

BUT artifacts can affect model fit and noise amplitude

NEED preprocessing to reduce artifacts



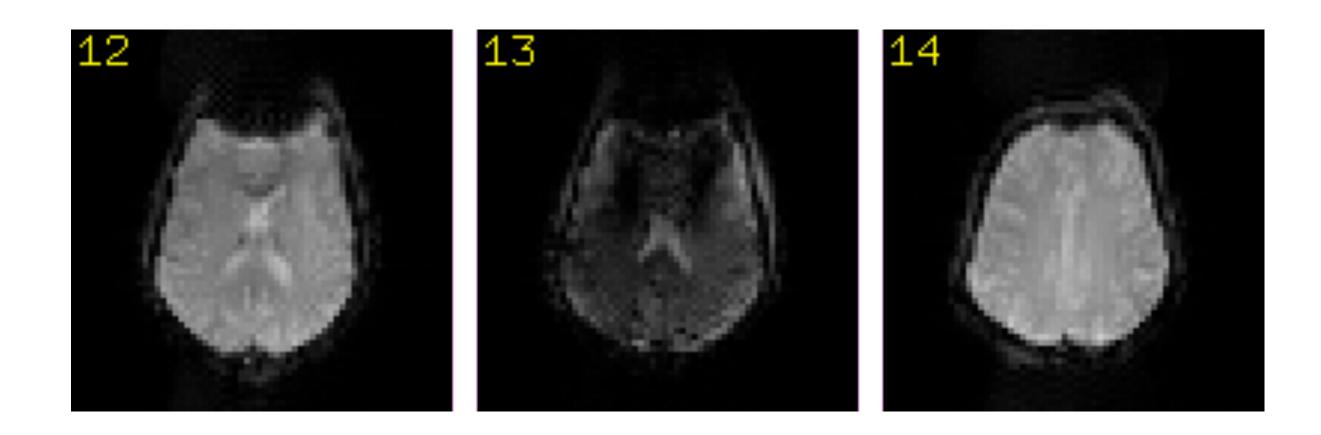


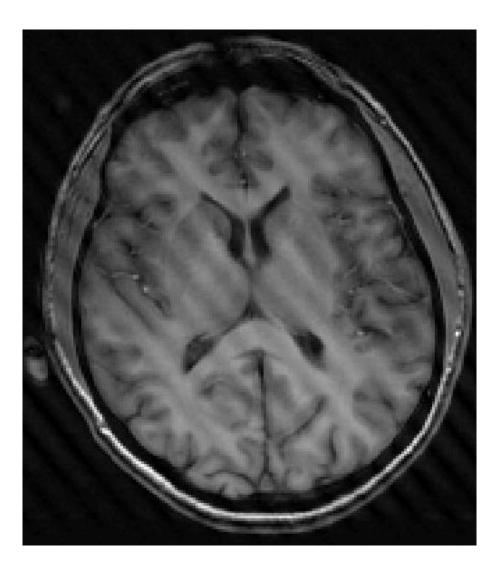
### Preprocessing



## Image reconstruction

- Convert k-space data to images using reconstruction algorithm
- Problematic data due to e.g., slice timing errors, RF spikes or RF interference









### Scanner artifact detection

#### LOOK AT YOUR DATA!





# Preprocessing overview

Conventional

Motion & distortion correction

High pass temporal filtering

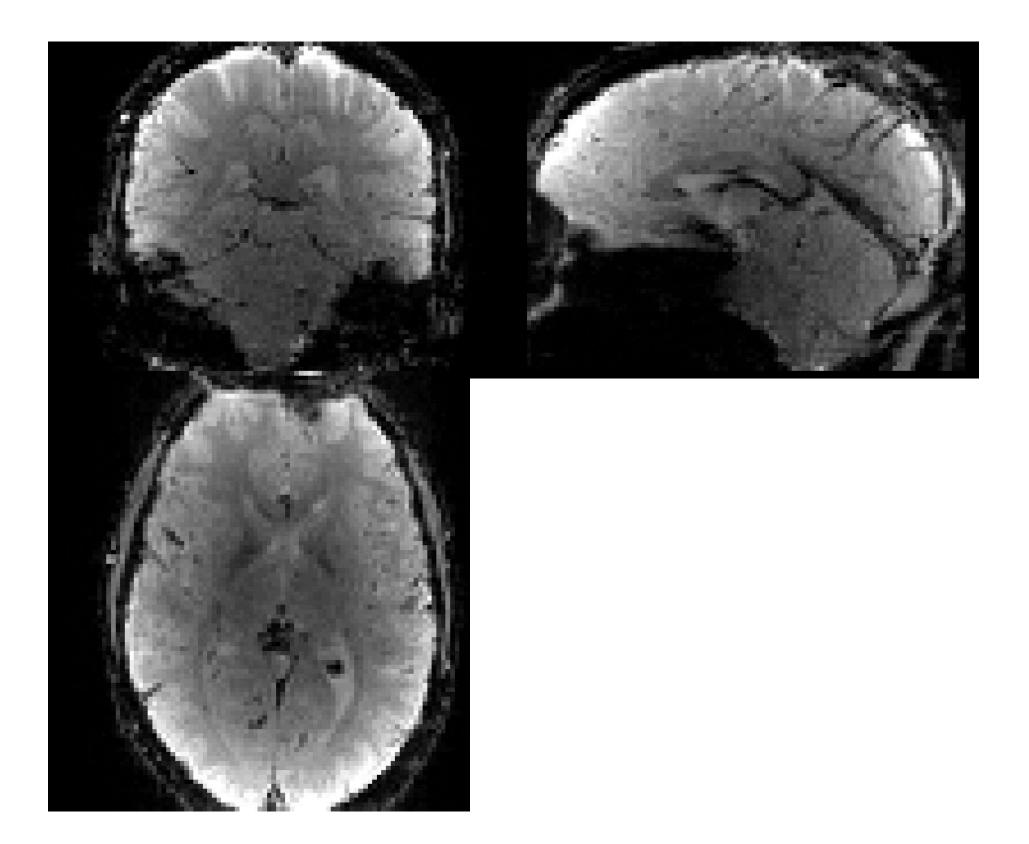
Registration

preprocessing steps							
		Slice timing correction					
		Spatial smoothing					

#### These same concepts are applied across all imaging modalities (task, rest, diffusion, ASL etc)



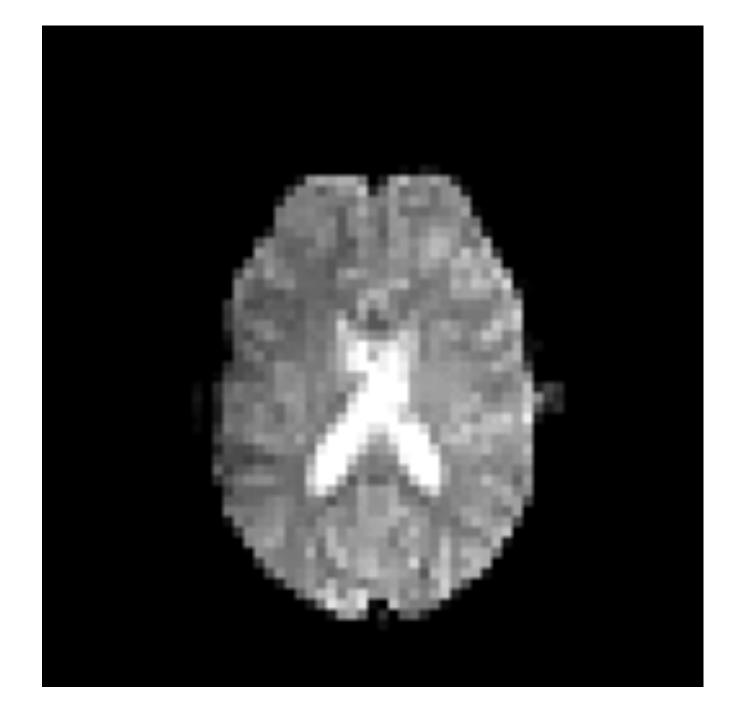
### Motion correction



- Biggest source of noise
- Correction is always necessary
- Linear correction may leave motion effects in data (e.g. partial volume)
  - Additional motion correction methods will be discussed later in the course



## **Motion correction**



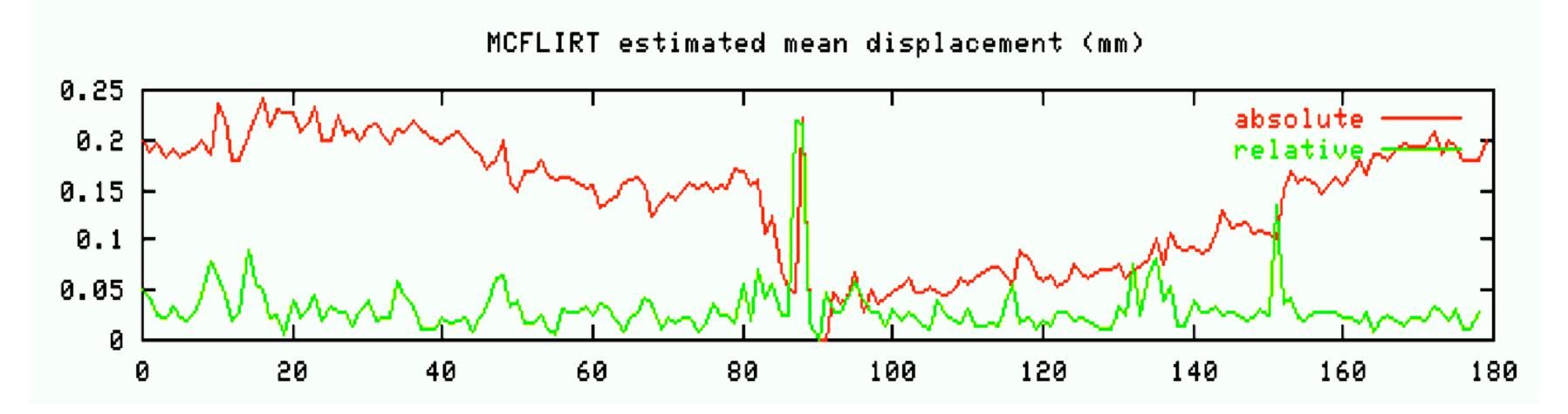
- Select reference timepoint = target
- Register each fMRI volume to target separately
- Use rigid body (6 DOF)

Uses rigid body registration



### **Motion correction**

- Results in 6 summary measures (rotations and translations)
  - Absolute = timepoint to target (shows jumps & drifts)
  - Relative = timepoint to next timepoints (shows jumps)
- Large jumps are more important than slow drifts (especially in relative motion plot)

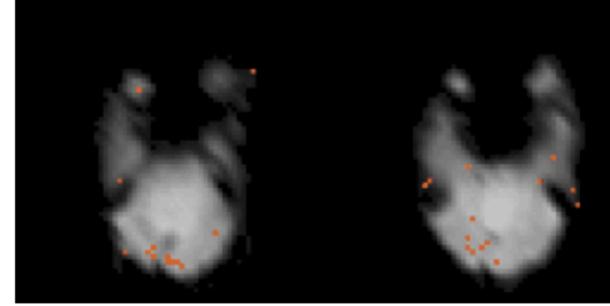




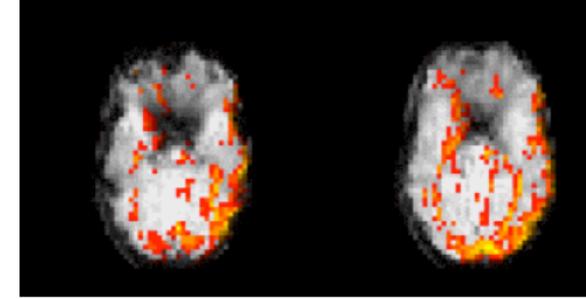


# Effect of motion correction

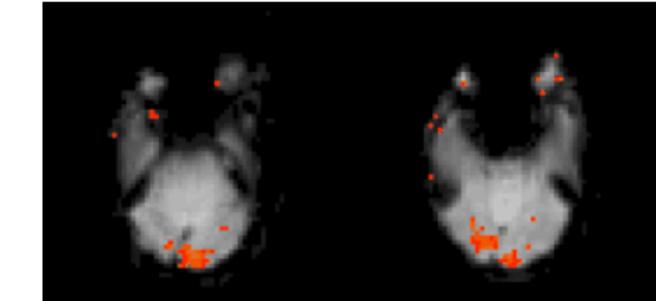
#### **Uncorrelated Motion**

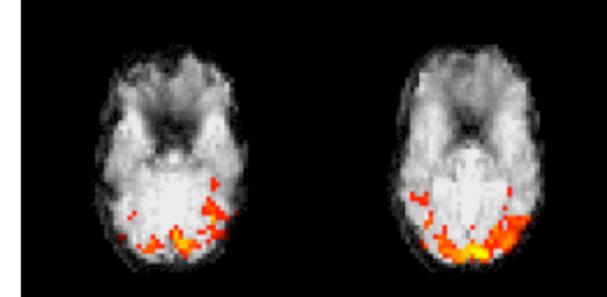


#### Stimulus Correlated Motion



#### Without MC





#### With MC



# Temporal filtering

#### Original BOLD data

# 

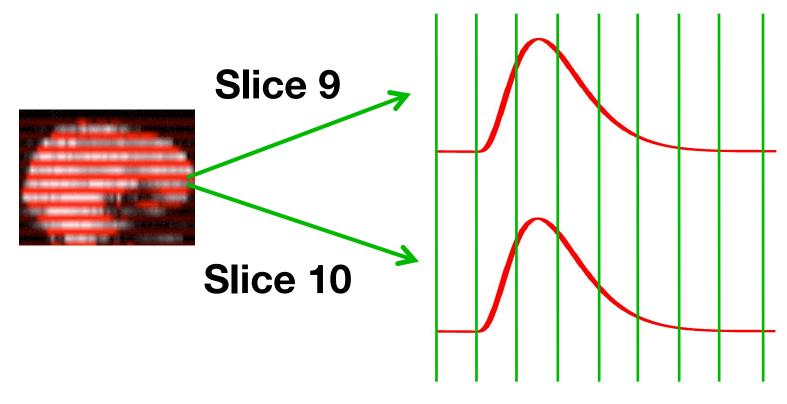
Highpass filtered data (>0.01 Hz)

- Correct for gradual drift in the data
- High-pass filtering recommended
- Need to choose cutoff frequency carefully
  - Lower frequency (i.e. longer period) than task frequencies of interest
  - Should be >90 seconds for autocorrelation estimation

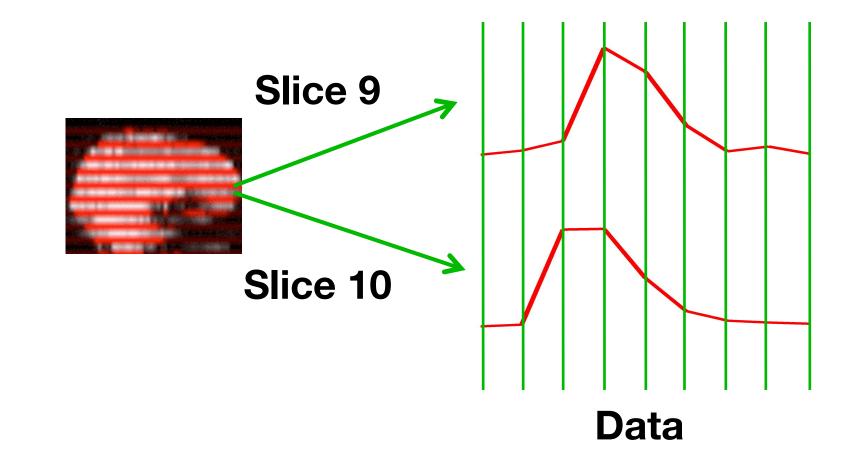




## Slice timing correction



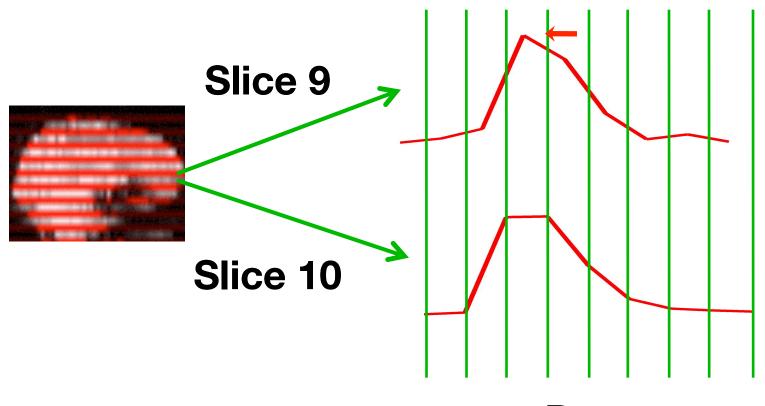
Model



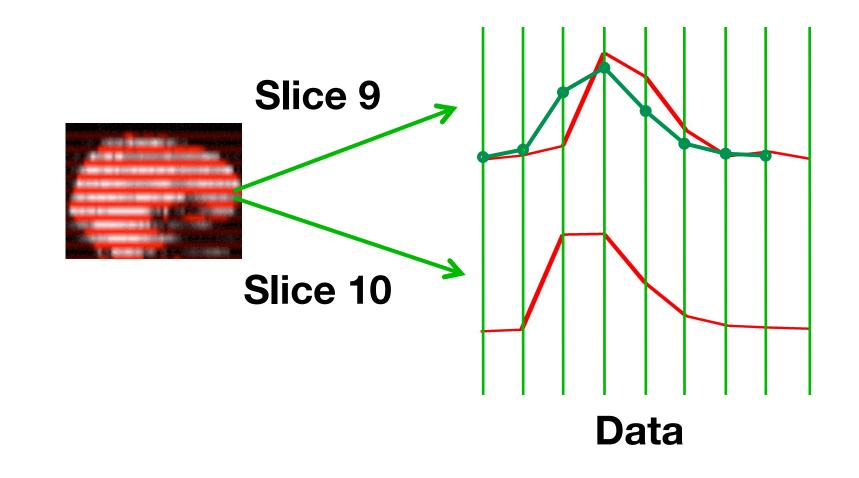
 Need to adjust for differences in acquisition time between different slices







Data





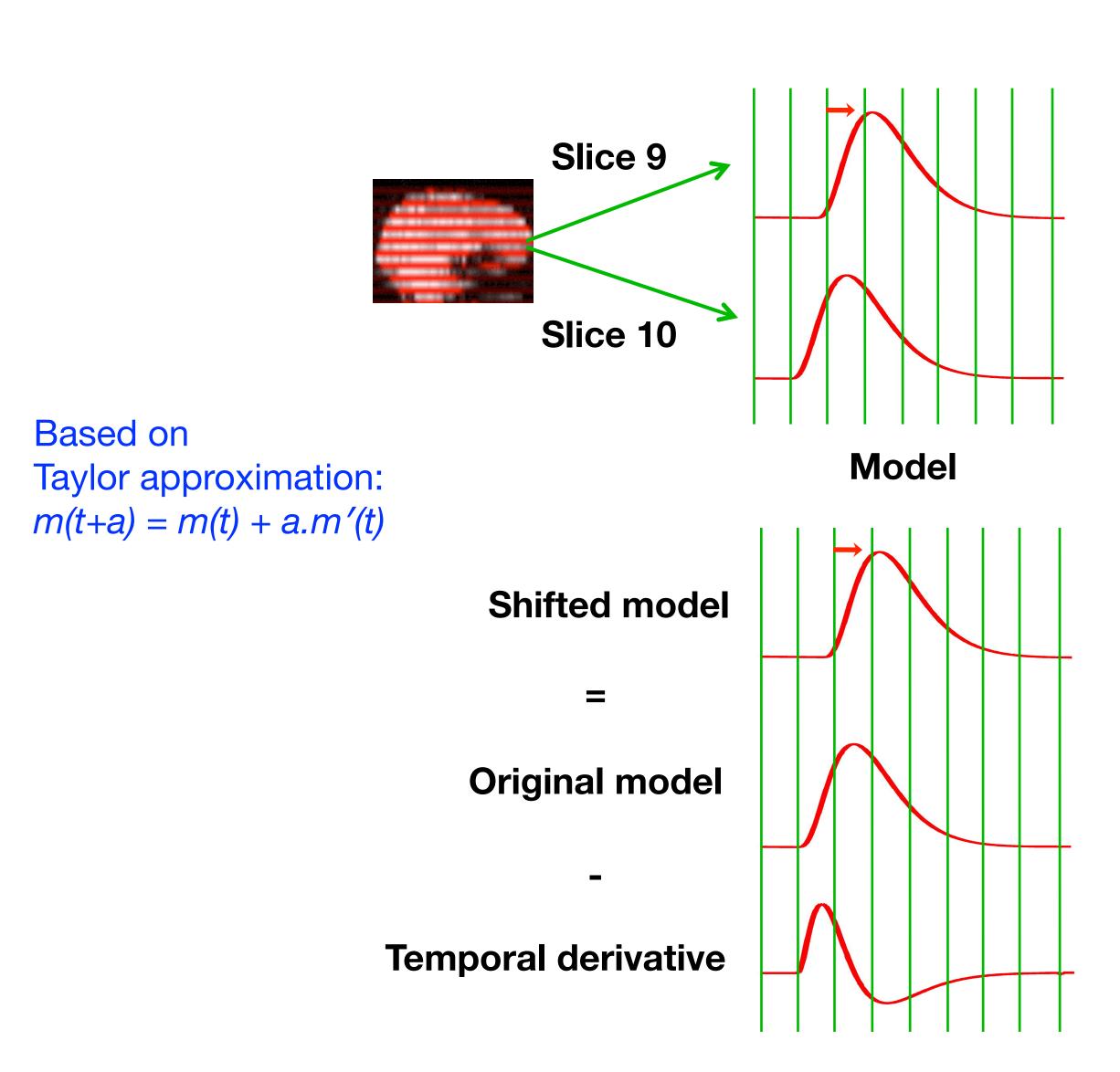
#### Slice timing correction

- Need to adjust for differences in acquisition time between different slices
- Can de with slice timing correction (i.e. shifting the data), but interpolation leads to degraded data





### Slice timing correction



- Need to adjust for differences in acquisition time between different slices
- Can de with slice timing correction (i.e. shifting the data), but interpolation leads to degraded data
- Better to shift the model, which we can do by including the temporal derivatives of EVs

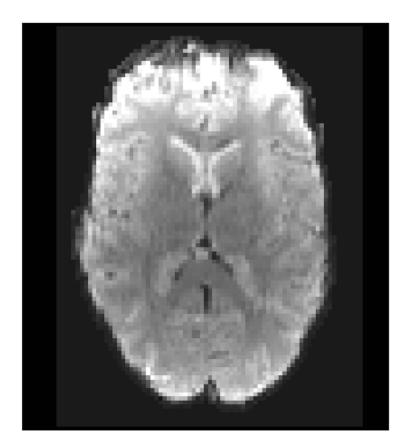




# Smoothing (spatial filtering)

FWHM

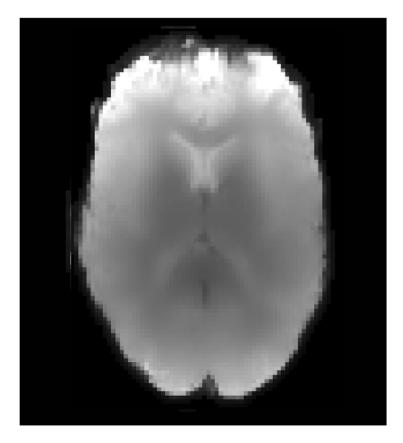
0.1	0.3	0.4	0.3	0.1			
0.3	0.6	0.8	0.6	0.3			
0.4	0.8	1.0	0.8	0.4			
0.3	0.6	0.8	0.6	0.3			
0.1	0.3	0.4	0.3	0.1			
FWHM							



No smoothing



5 mm smoothing



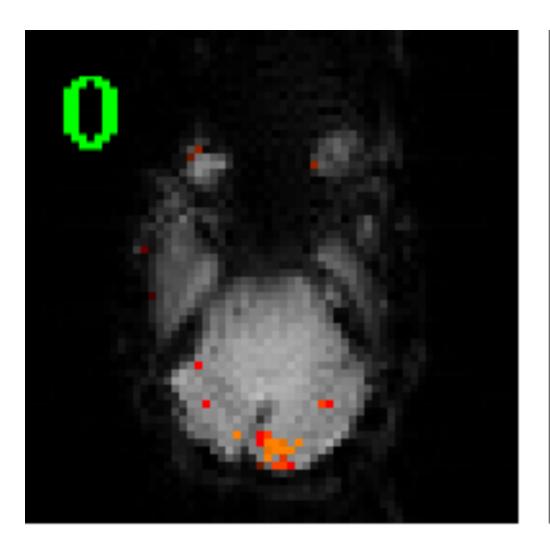
10 mm smoothing

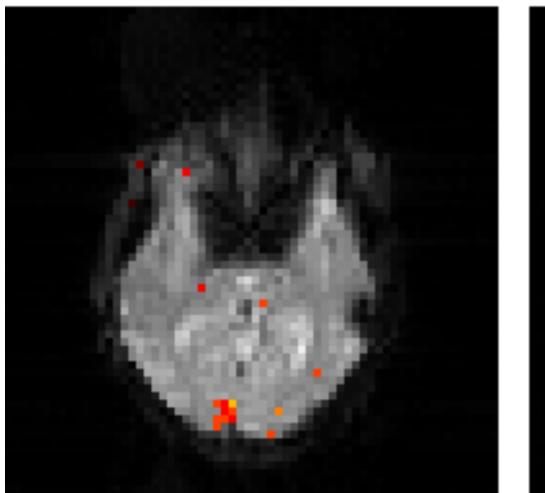
- Increase signal to noise ratio by lacksquareblurring the image
  - Averaging leads to less noisy voxel values
  - Needed for some stats (GRF)
- Keep in mind:
  - Size of expected activation
  - **Desired spatial specificity**
  - May not need to do any

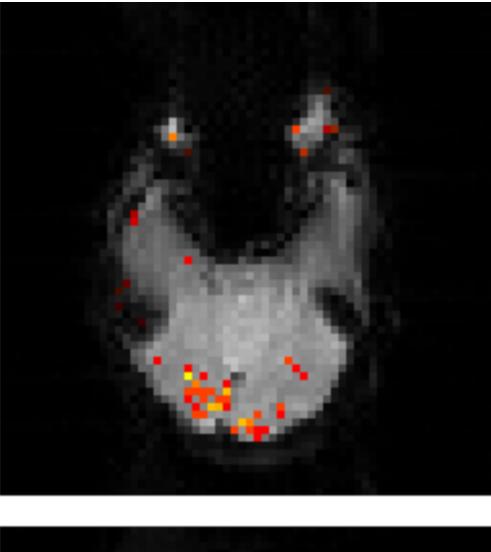


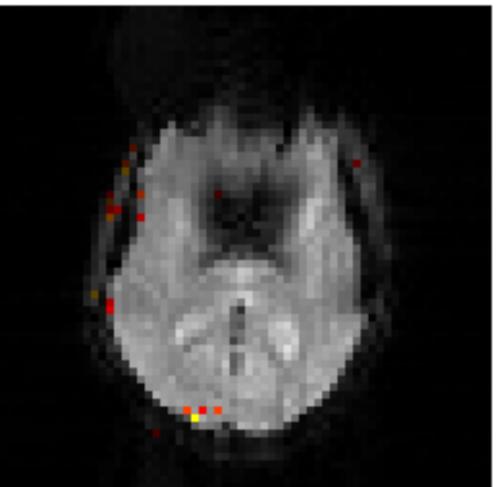


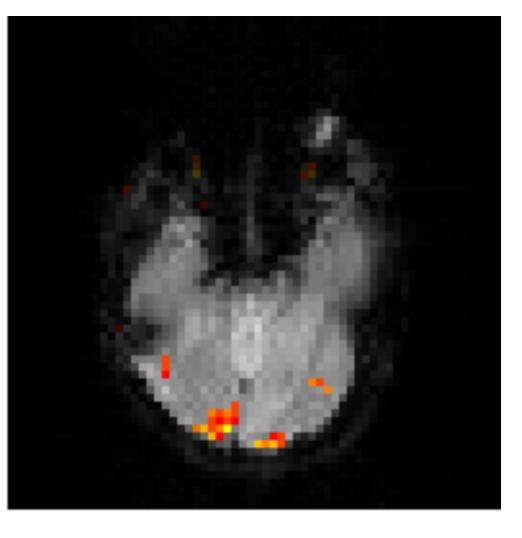
### Effect of smoothing

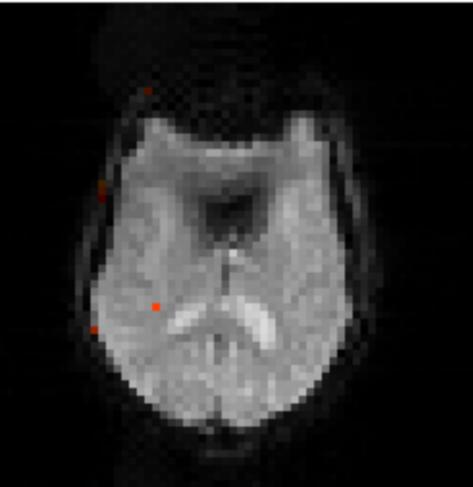






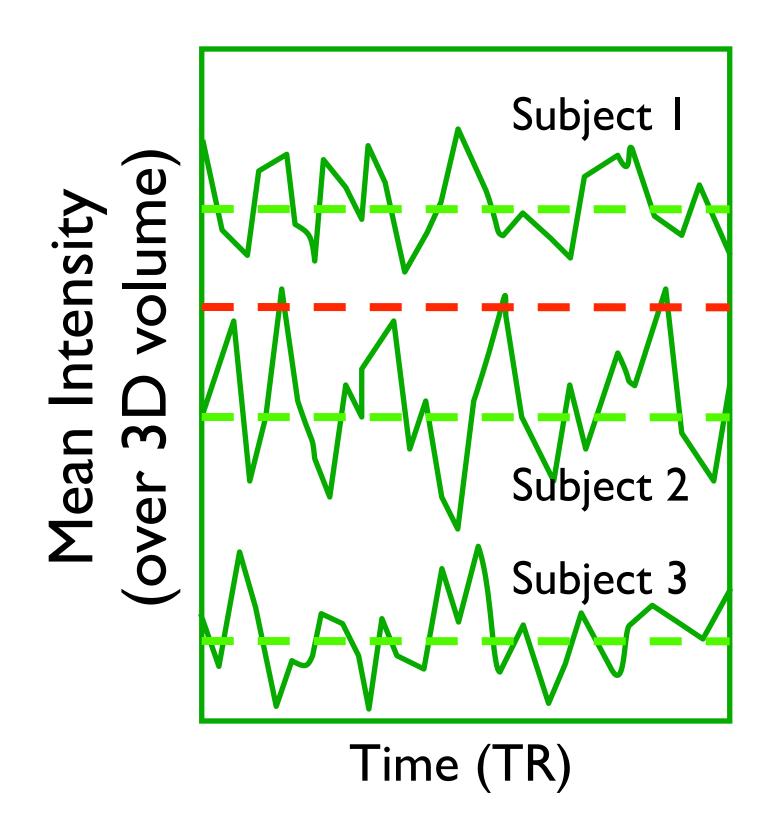








#### Global Intensity Normalisation



- Need to remove uninterested offsets between subjects and sessions
- Scale (i.e. multiply) each 4D dataset by a single value
- Automatically done in all software packages
- Not the same as global signal regression!



# Summary of preprocessing

Reconstruction

Motion Correction

Slice Timing

**Spatial Smoothing** 

**Temporal Filtering** 

Intensity Normalisation

- Create image and remove gross artefacts
- Get consistent anatomical coordinates (always do this)
- Get consistent acquisition timing (use temporal derivative instead)
- Improve SNR & validate GRF
- Highpass: Remove *slow* drifts Lowpass: Avoid for autocorr est.
- 4D: Keeps overall signal mean constant across sessions



#### Break Time!



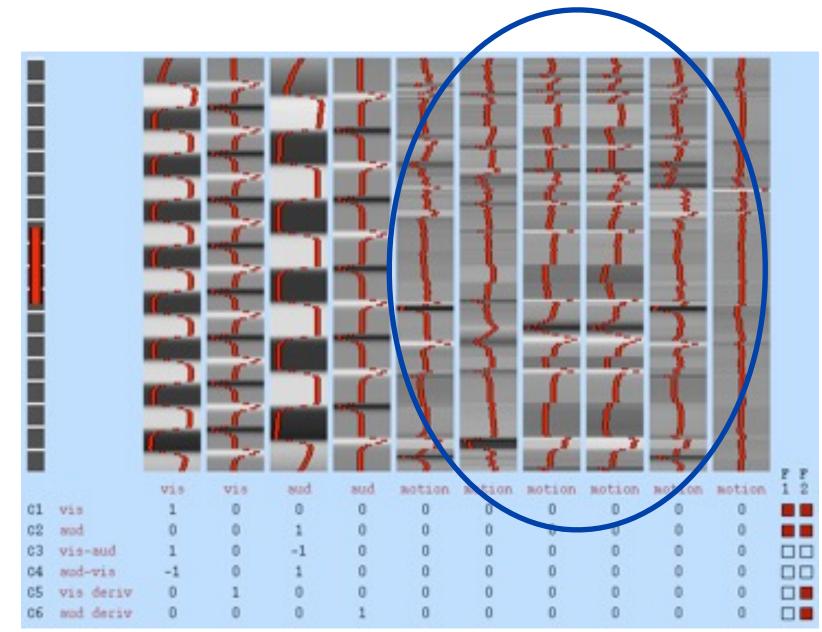


# **Motion Parameter Confounds**

GLM as confounds - simple button in FEAT

- Removes any correlated signals (since they are confounds)
- Assumes a linear relationship between motion parameters and intensity of motion artefact
- Assumes that MCFLIRT estimation is accurate
- Problematic if motion parameters and EVs of interest are highly correlated (stimulus-correlated motion)

Add the 6 parameters (rotations and translations) as measured by MCFLIRT to the

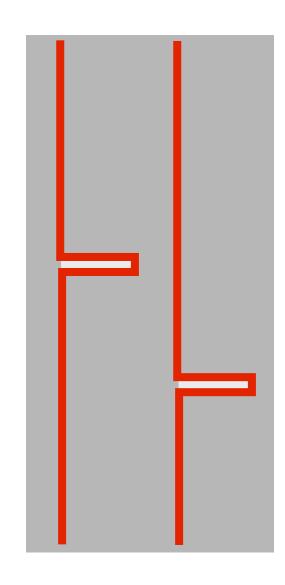


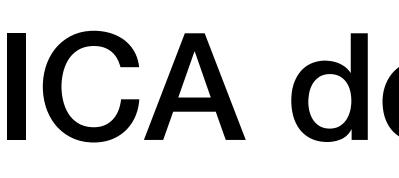


## **Outlier Timepoint Detection**

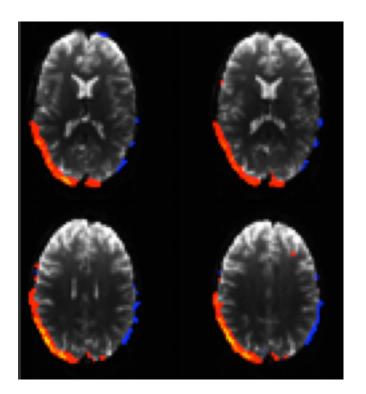
- Also known as scrubbing or volume censoring
- Removes **all** influence of the timepoints
- Uses one extra confound regressor per outlier timepoint
- fsl\_motion\_outliers creates confound matrix for GLM
- Can cope with very extreme motion effects
- But leaves other timepoints uncorrected

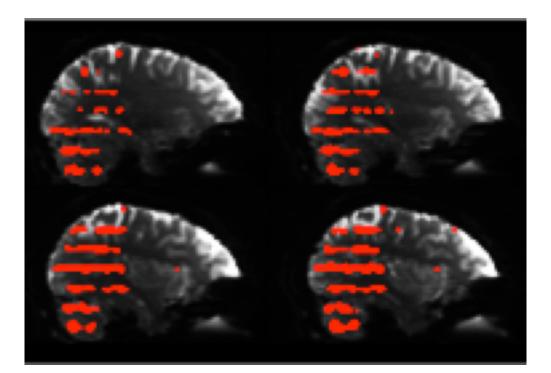
Confound matrix with 2 outlier timepoints



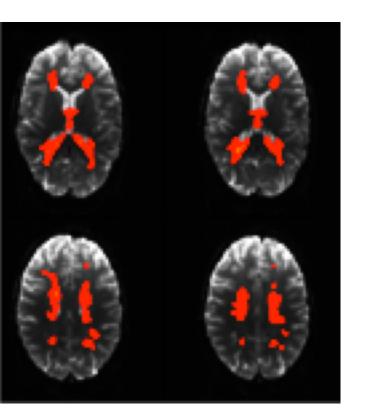


- Use ICA to identify noise comports
- More on this during the ICA lecture



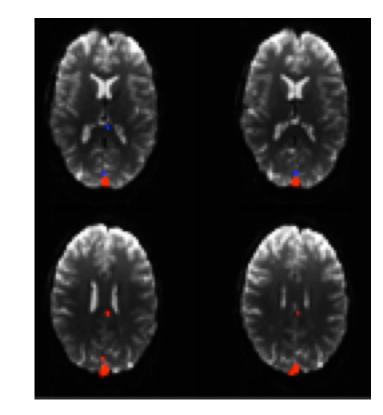


#### Classic motion



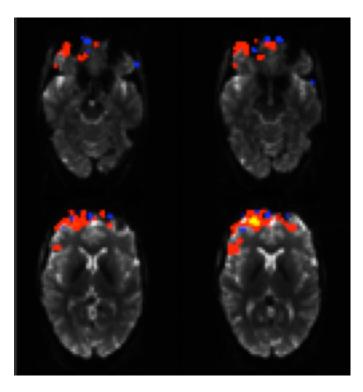
#### White matter



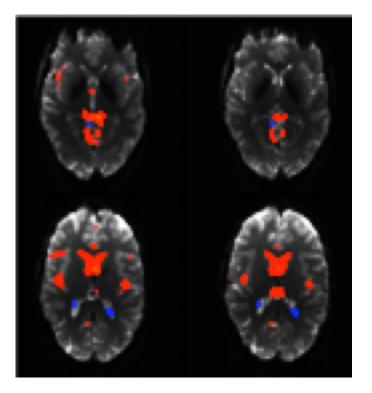


Sagittal sinus

### ICA denoising



#### Multiband motion Susceptibility motion

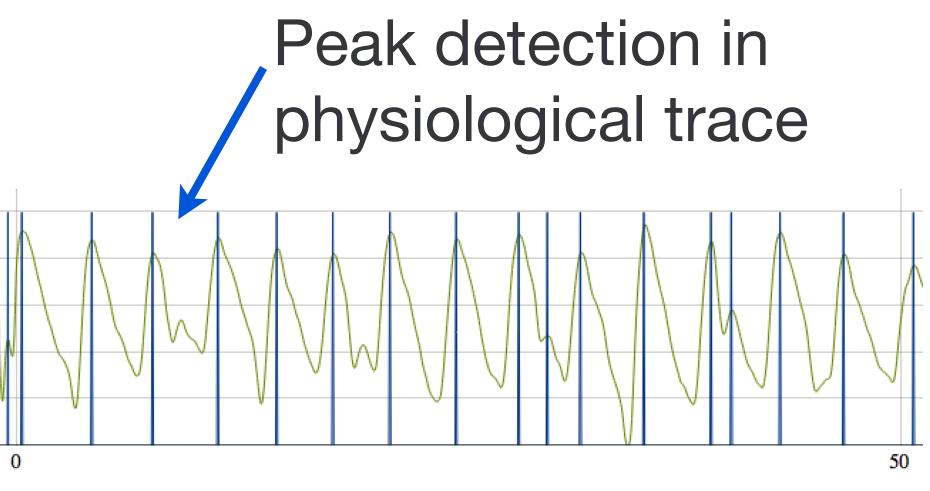


Cardiac/CSF



# **Physiological Noise Regression**





Very important for high-risk brain regions such as the brainstem

**PNM GUI creates a** set of files suitable for use as *Voxelwise* Confounds in FEAT

