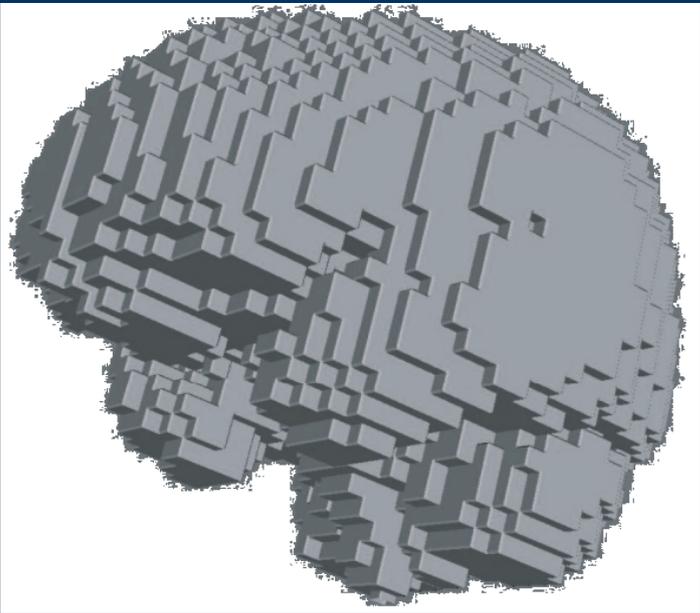
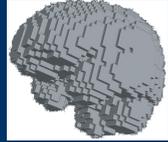
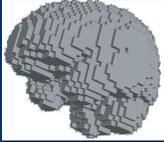
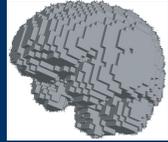
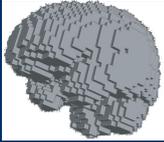


FMRI data preprocessing

Martin Monti
UCLA Department of Psychology

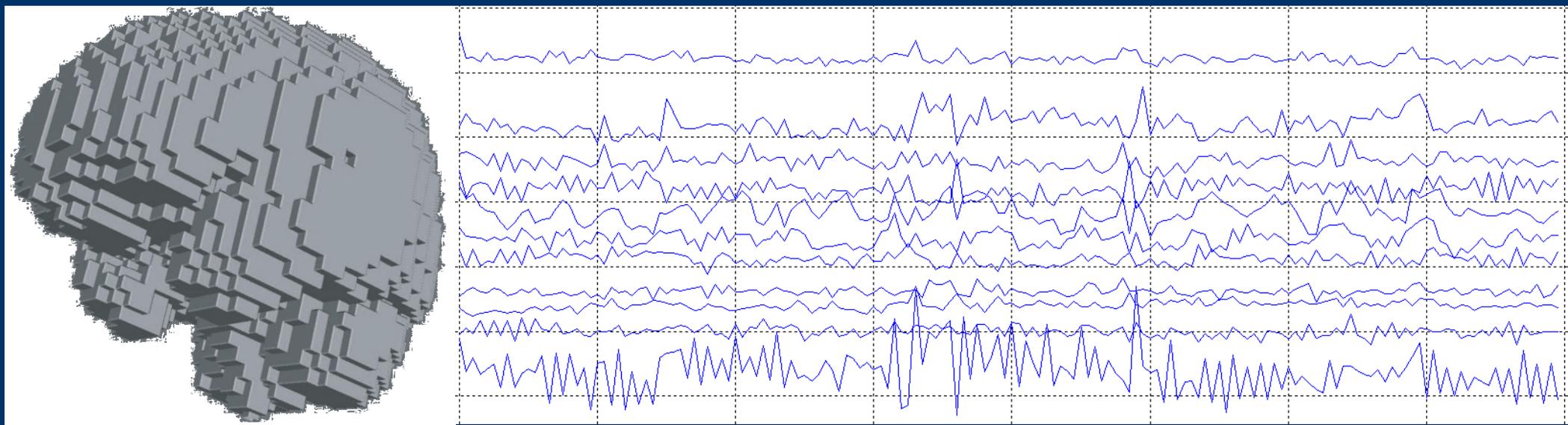


Typical dataset



Typical dataset

Run



Typical fMRI analysis sequence

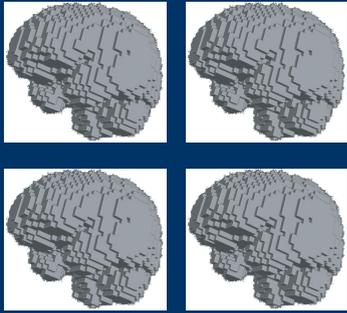
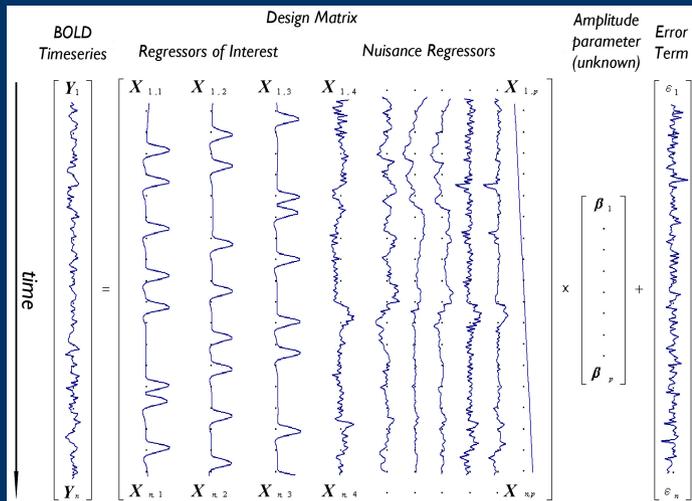
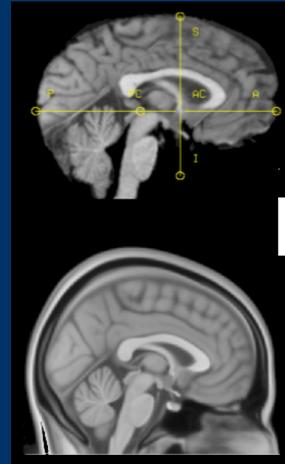


Image Pre-processing

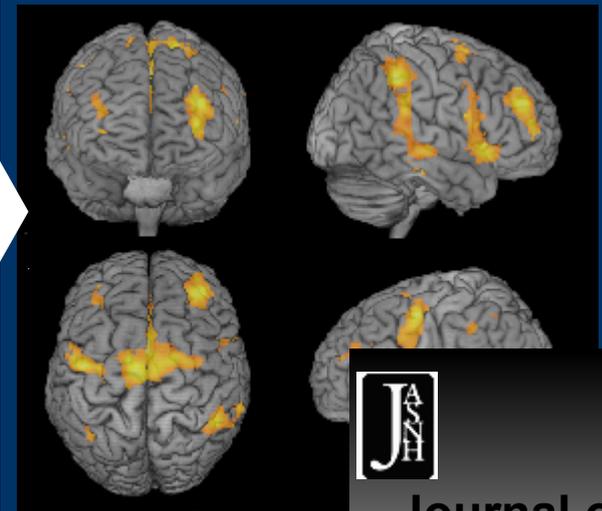
Single Subject Analysis



Normalization



Group Analysis



Journal of
Articles in
Support of
The Null
Hypothesis

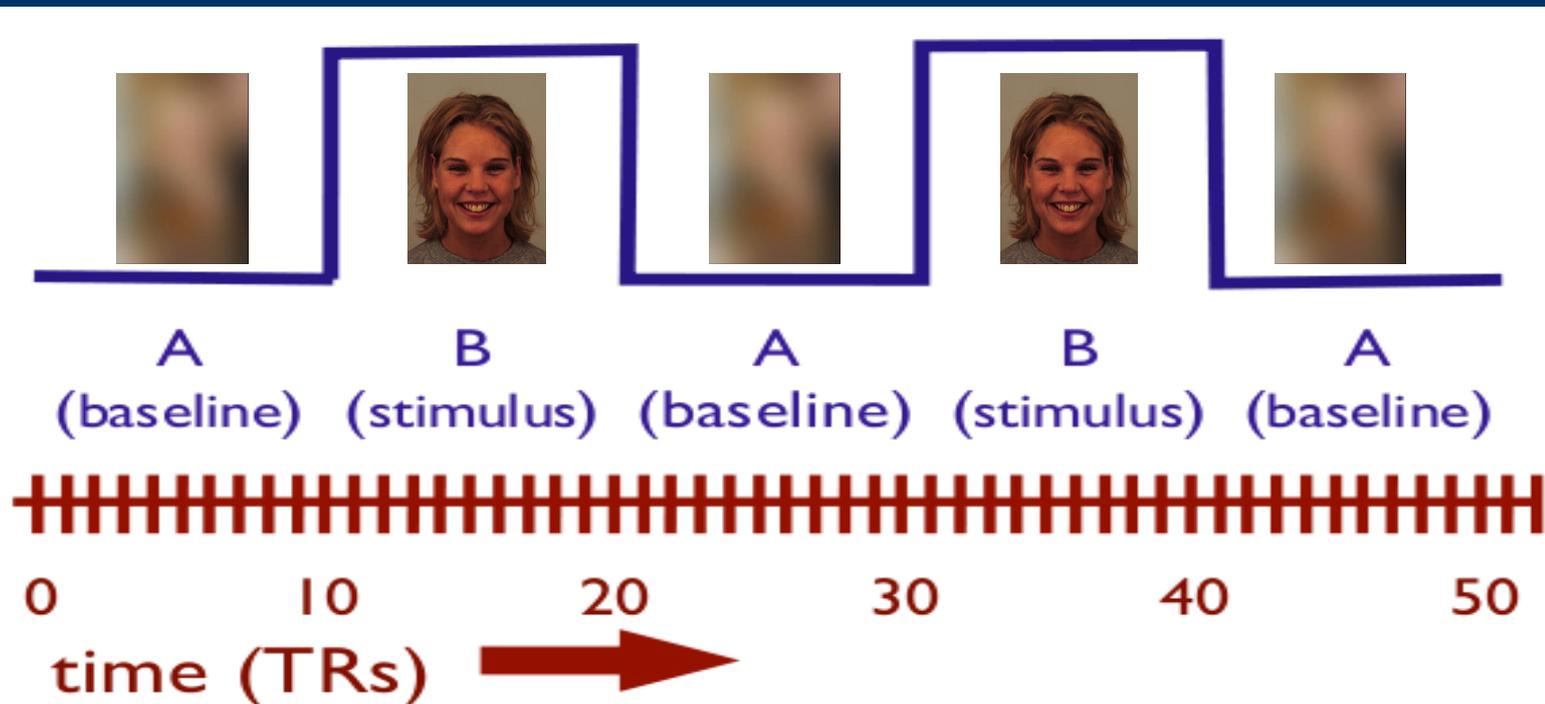
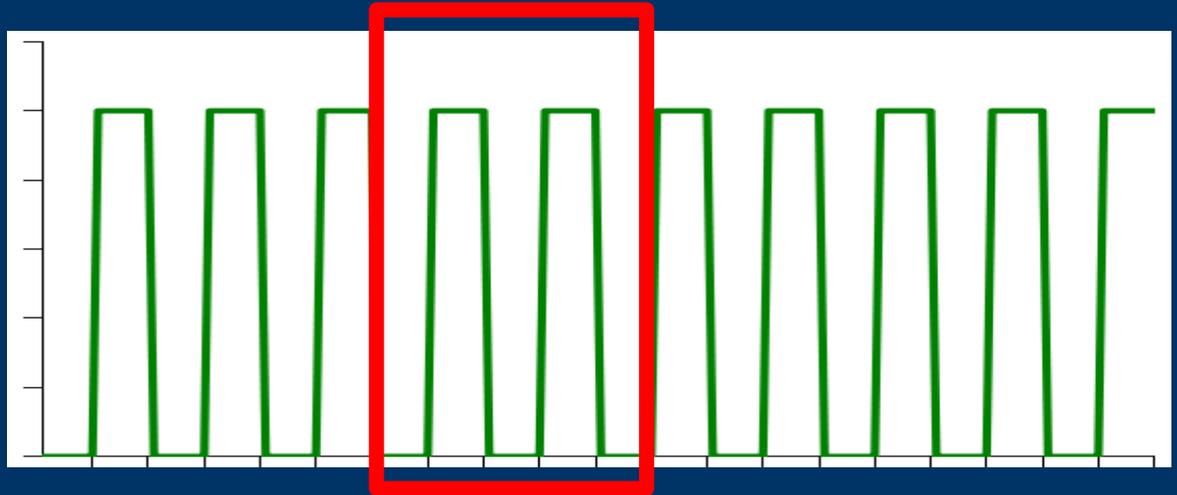
Preprocessing: what/why?

Preprocessing is a series of data transformations (“data conditioning”) aimed at reducing sources of noise in order to:

- 1) Increasing sensitivity of analysis (SNR)
 - 2) Ensuring validity of the statistical model
-
-

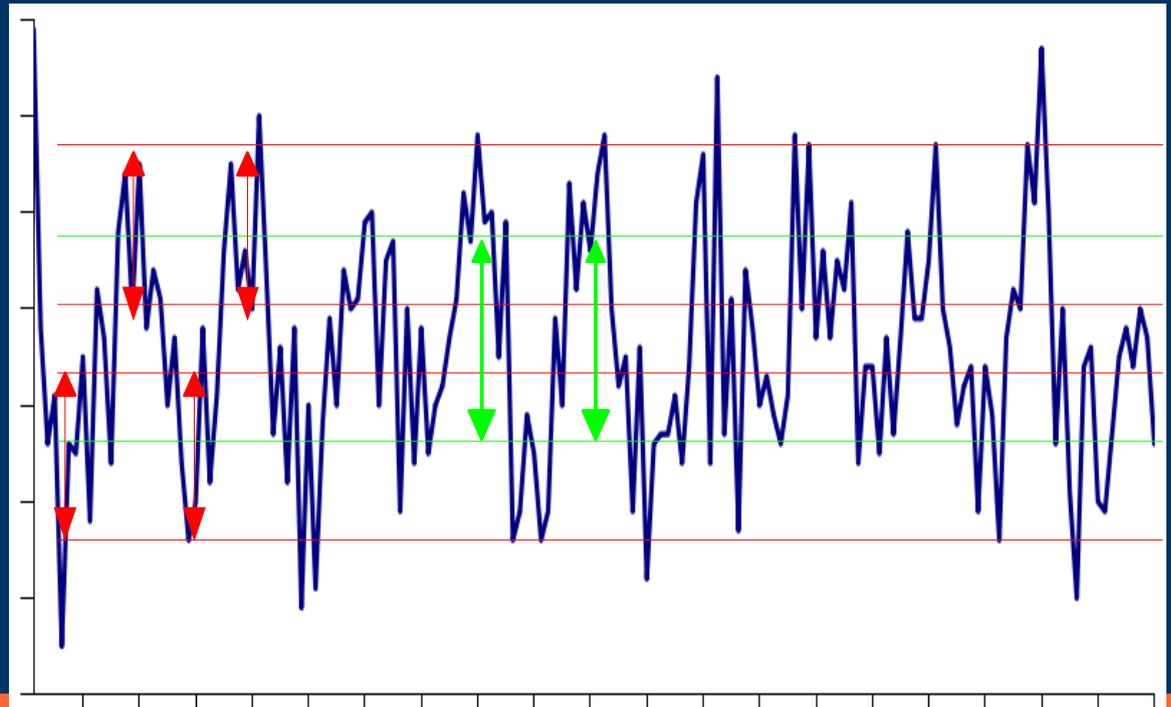
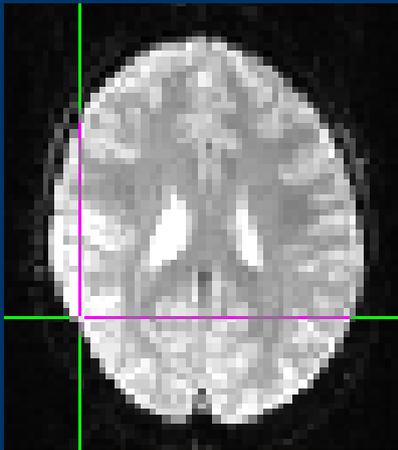
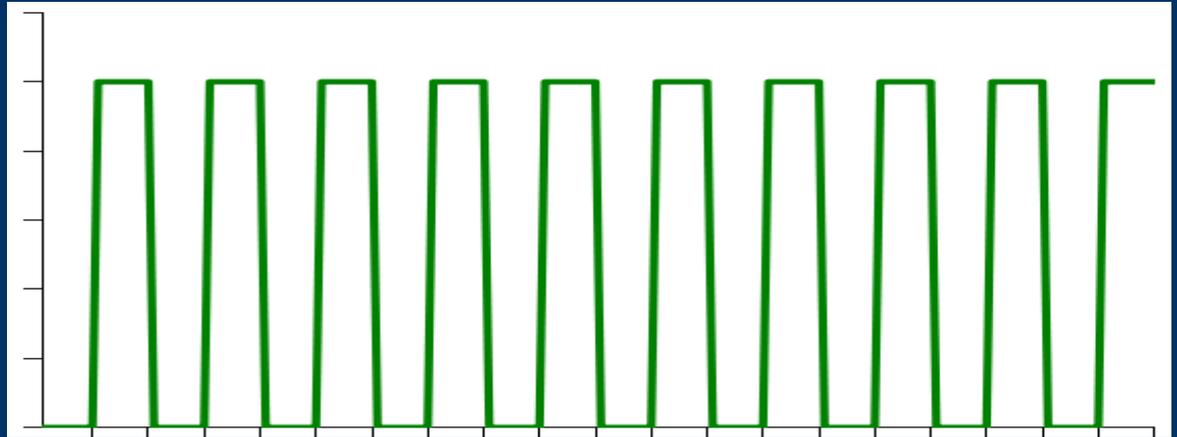
Sample Experiment: SNR

TR = 2s
Vols = 160
10 AB Cycles
Cycle = 8A + 8B



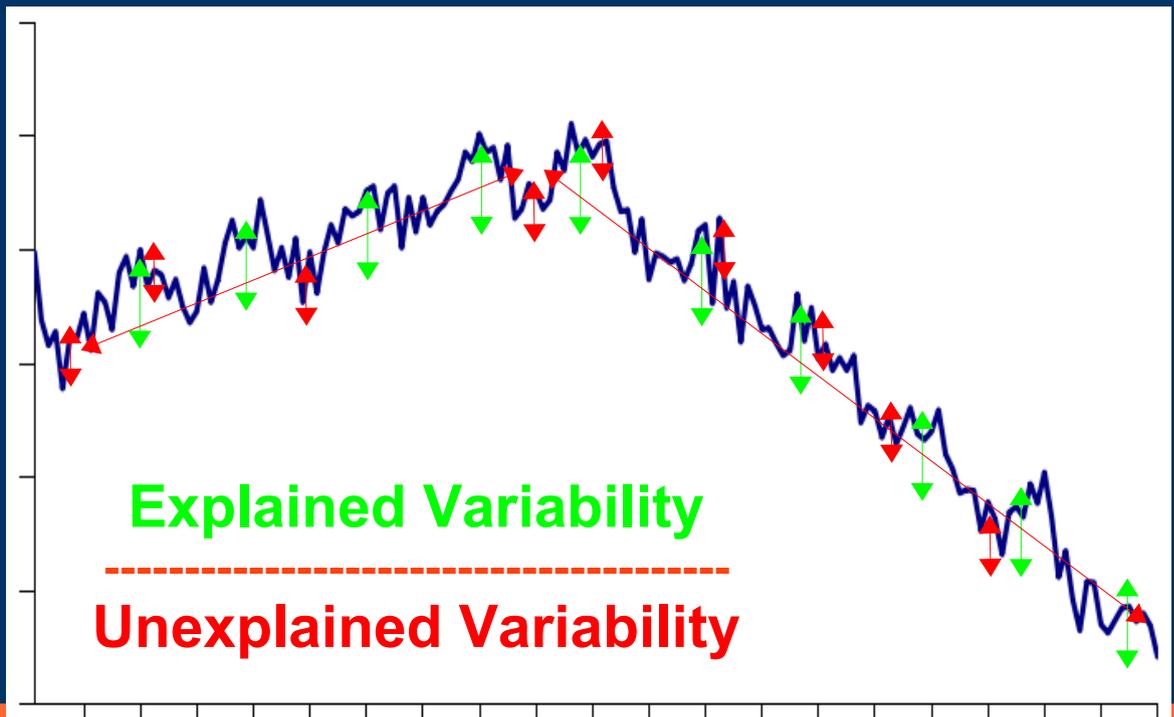
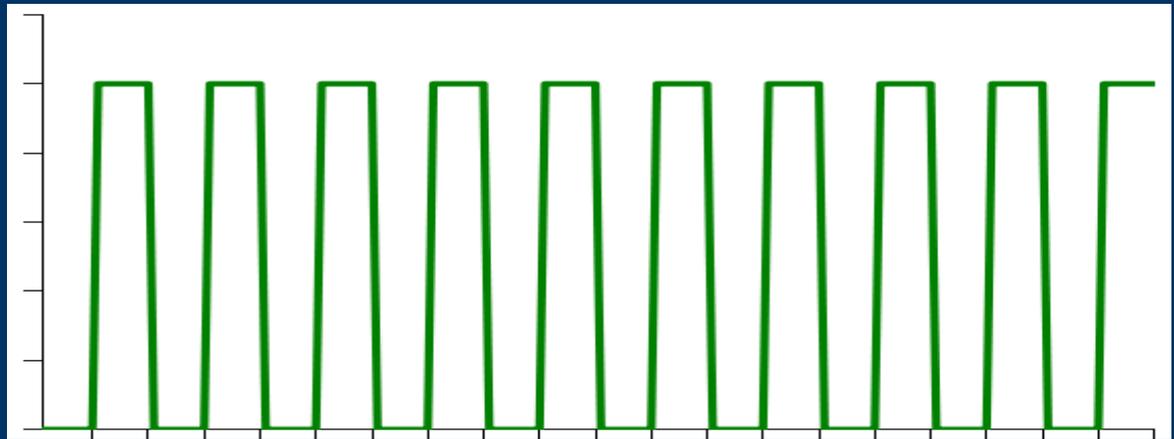
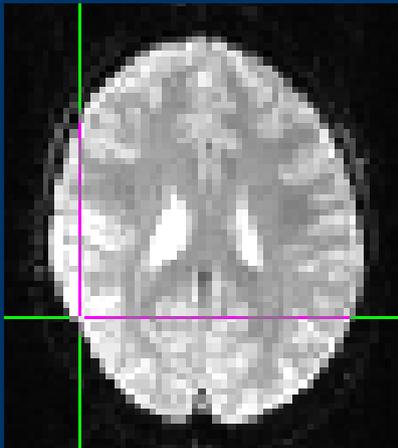
Sample Experiment: SNR

TR = 2s
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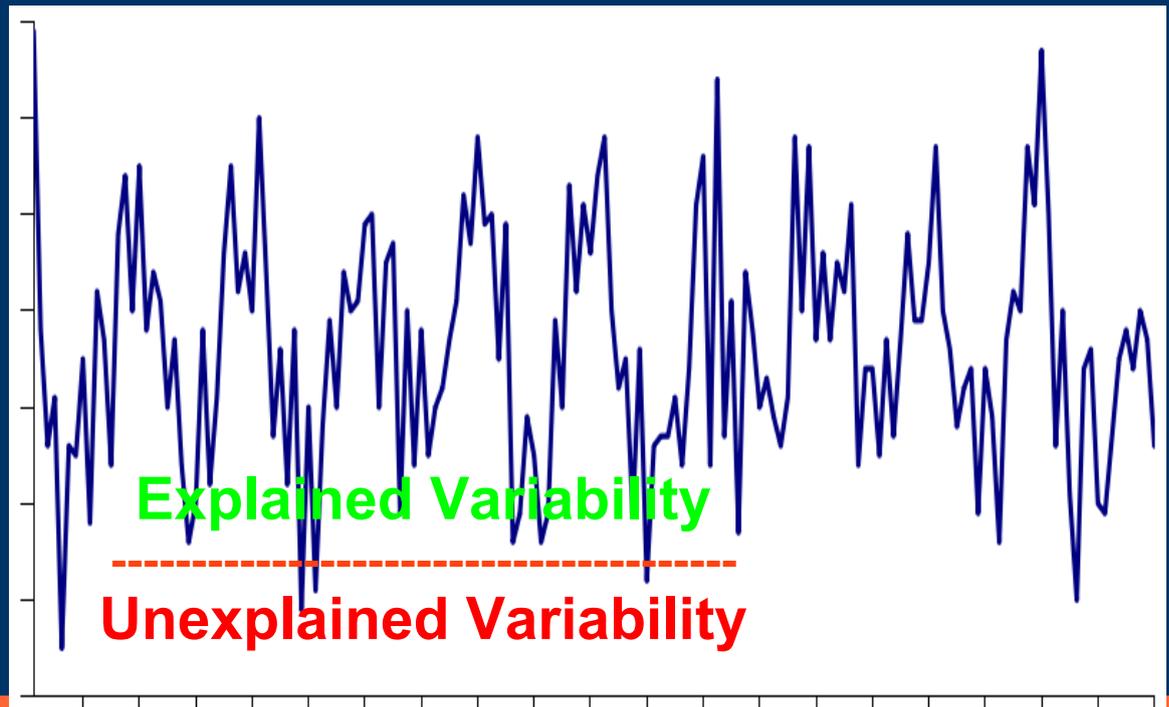
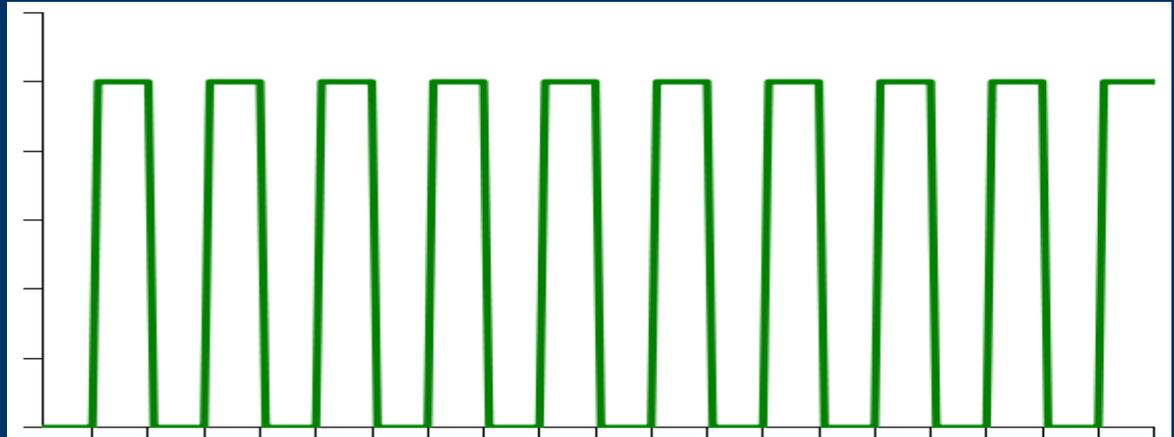
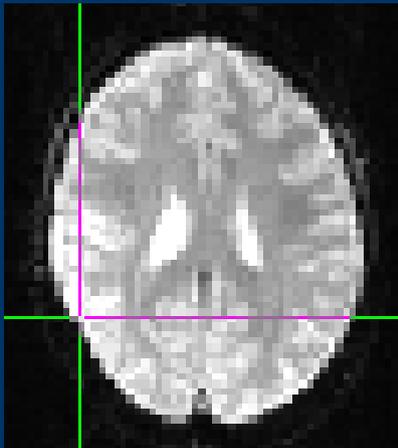
Sample Experiment: SNR

TR = 2s
Vols = 160
10 AB Cycles
Cycle = 8A + 8B



Sample Experiment: SNR

TR = 2s
Vols = 160
10 AB Cycles
Cycle = 8A + 8B

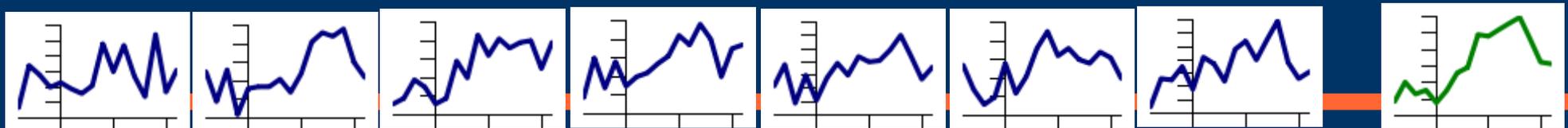
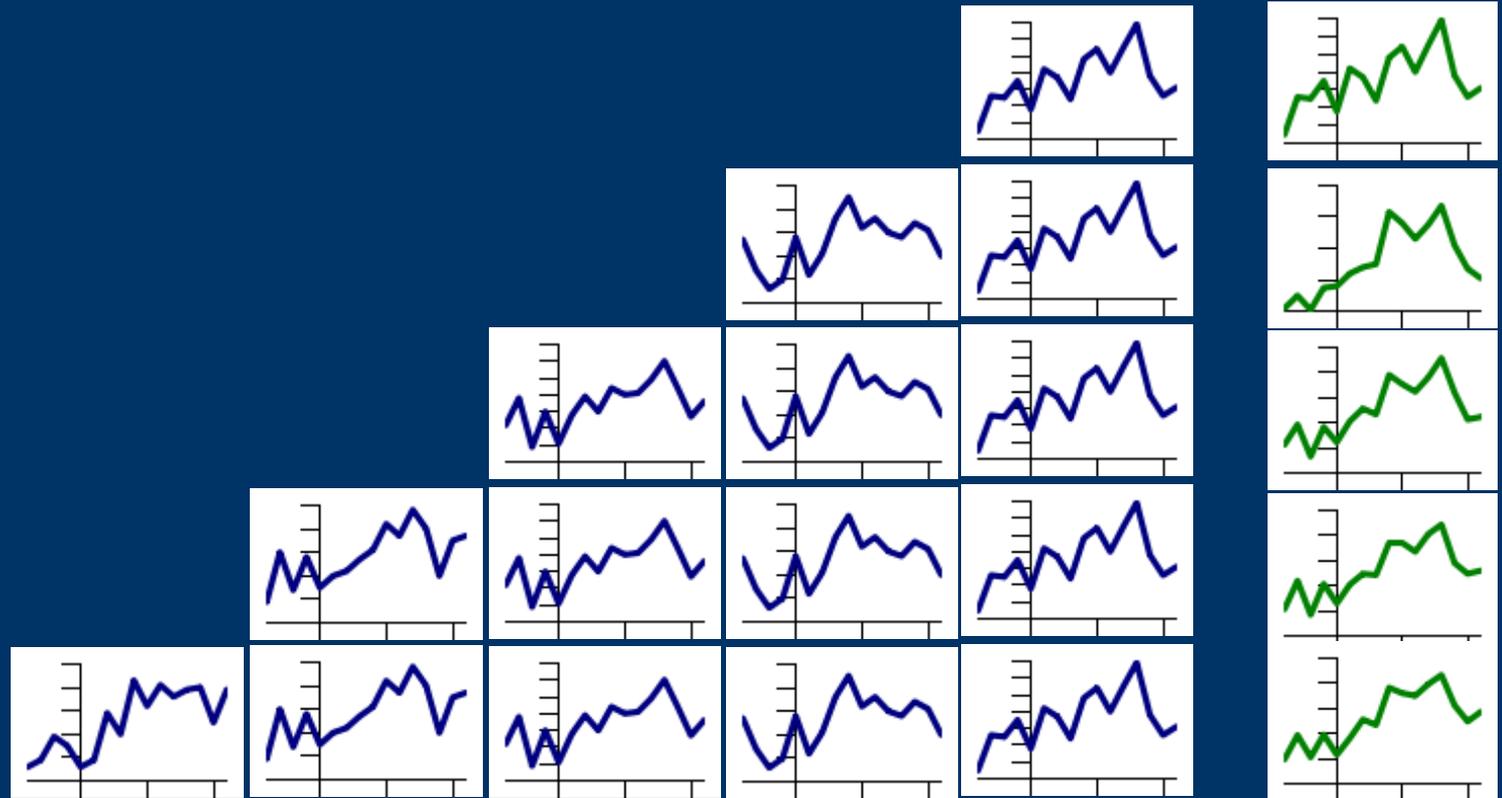


Noise in fMRI

- Hardware & acquisition related
 - Thermal noise (intrinsic noise)
 - System noise (scanner drift)
 - Field inhomogeneities
 - Slice acquisition timing
 - ...
 - Subject related
 - Oscillatory physiological noise (heartbeat, respiration)
 - Head motion
 - Psychological (alertness, learning)
 - White noise
-
-

Coping with Gaussian noise

AVERAGE



Noise in fMRI

- Hardware & acquisition related
 - Thermal noise (intrinsic noise)
 - System noise (scanner drift)
 - Field inhomogeneities
 - Slice acquisition timing
 - ...
 - Subject related
 - Oscillatory physiological noise (heartbeat, respiration)
 - Head motion
 - Psychological (alertness, learning)
 - White noise
-
-

The General Linear Model (GLM)

$$y = X \times \beta + \varepsilon$$

fMRI Signal = Design Matrix × Parameter + Residuals

“our data” = *“what we CAN explain”* × *“how much of it we CAN explain”* + *“what we CANNOT explain”*

Preprocessing: what/why?

Preprocessing is a series of data transformations (“data conditioning”) aimed at reducing sources of noise in order to:

- 1) Increasing sensitivity of analysis (SNR)
 - 2) Ensuring validity of the statistical model
-
-

Preprocessing

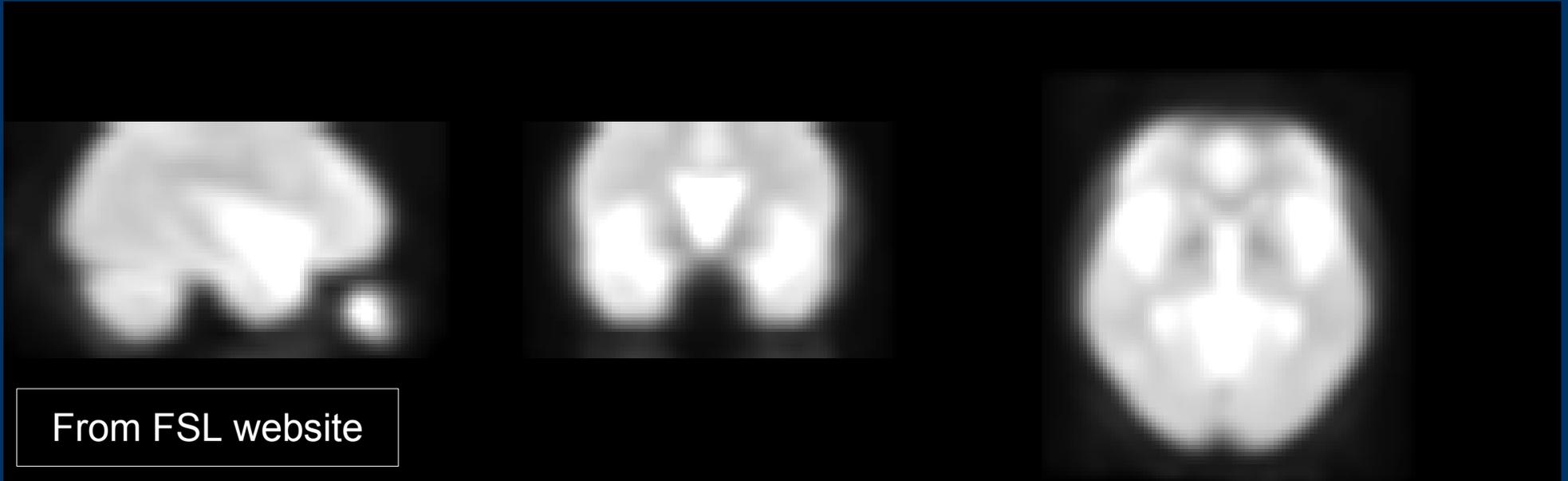
- i. Motion correction
 - ii. Slice timing correction
 - iii. Spatial filtering
 - iv. Temporal filtering
 - v. Intensity normalization
 - vi. Field distortion correction (unwarping)
-
-

Preprocessing

- i. Motion correction
- ii. Slice timing correction
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- iv. Temporal filtering
- v. Intensity normalization
- vi. Field distortion correction (unwarping)



Subject Motion

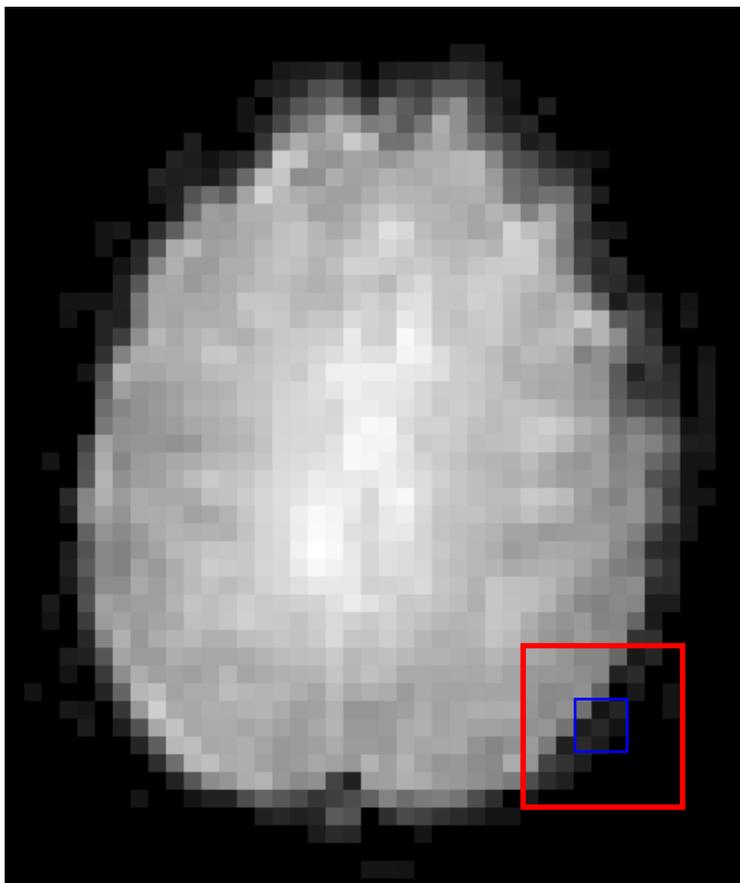


Motion within a time-series can have several unwanted consequences:

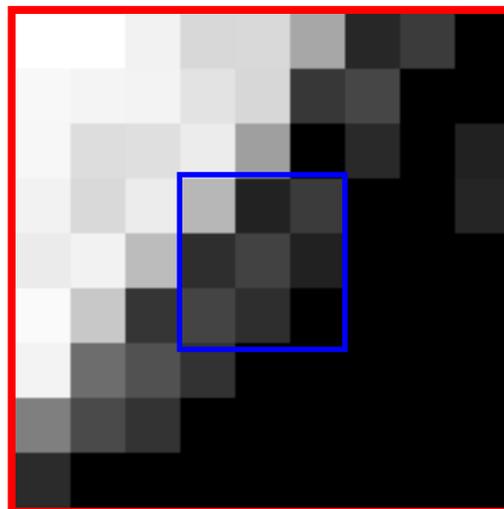
- Motion can produce signal changes of a greater magnitude than the BOLD signal
 - Lose the correspondence between a voxel and anatomical location
-
-

Subject Motion

A

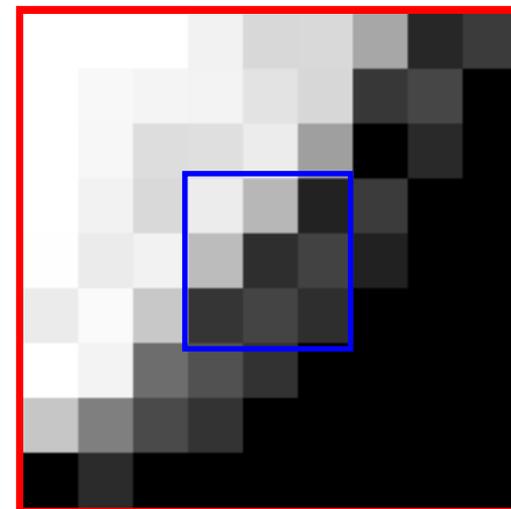


B



507	89	154
119	171	83
179	117	53

C

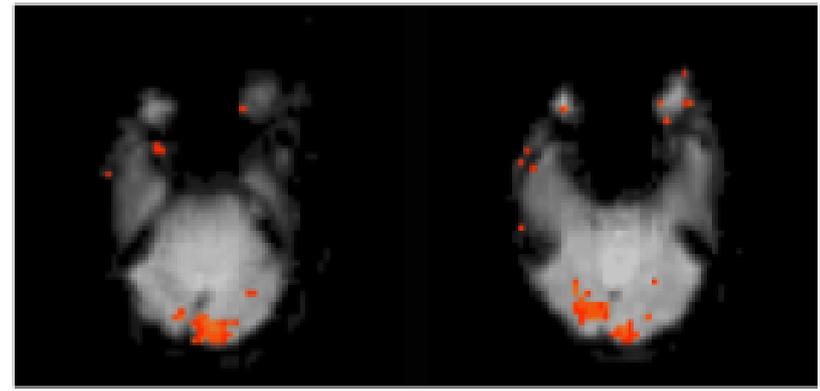
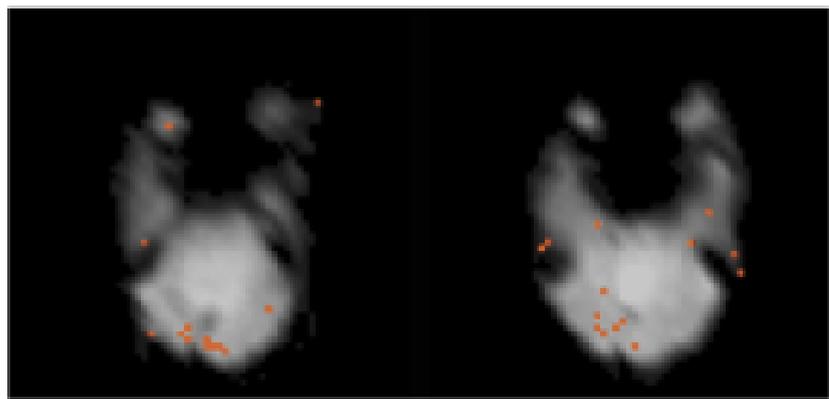


663	507	89
520	119	171
137	179	117

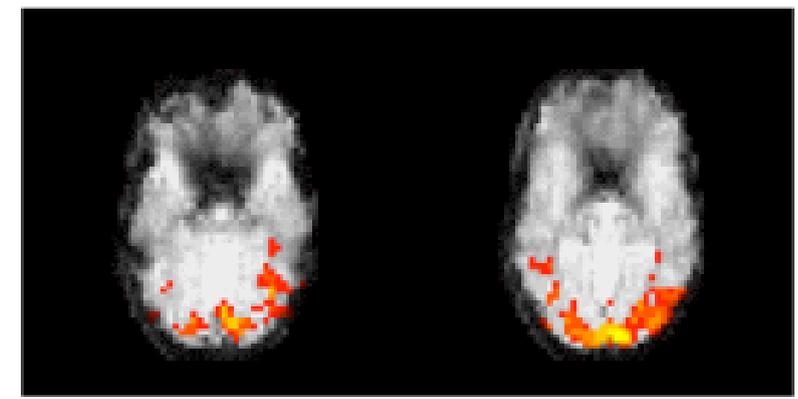
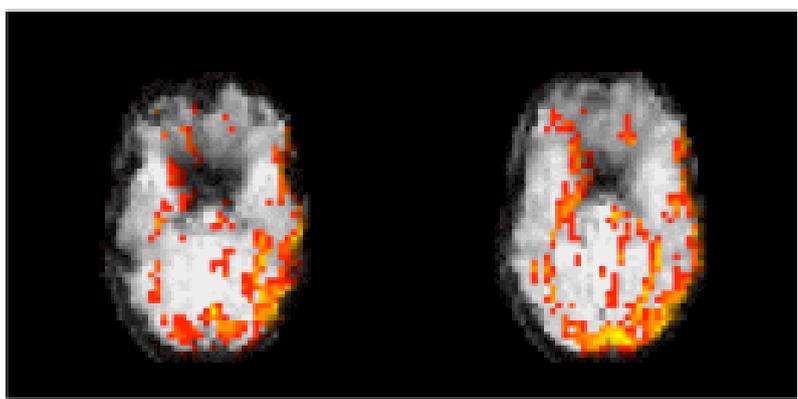
Huettel et al. Functional Magnetic Resonance Imaging



Effect of Motion Correction



Stimulus Correlated Motion



Without MC

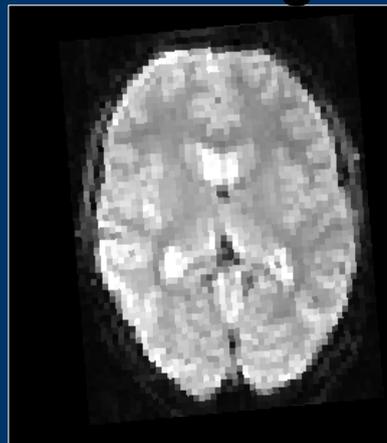
With MC

Motion Correction

Reference



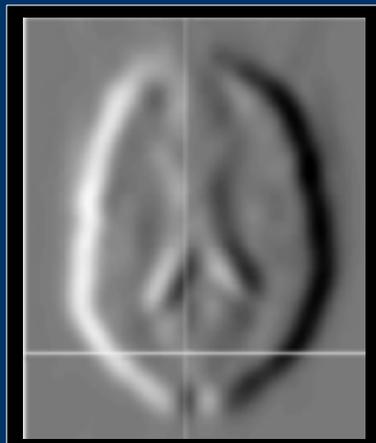
I th Image



-

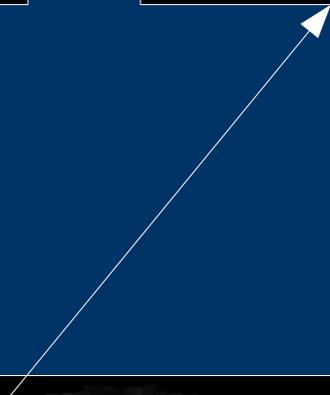
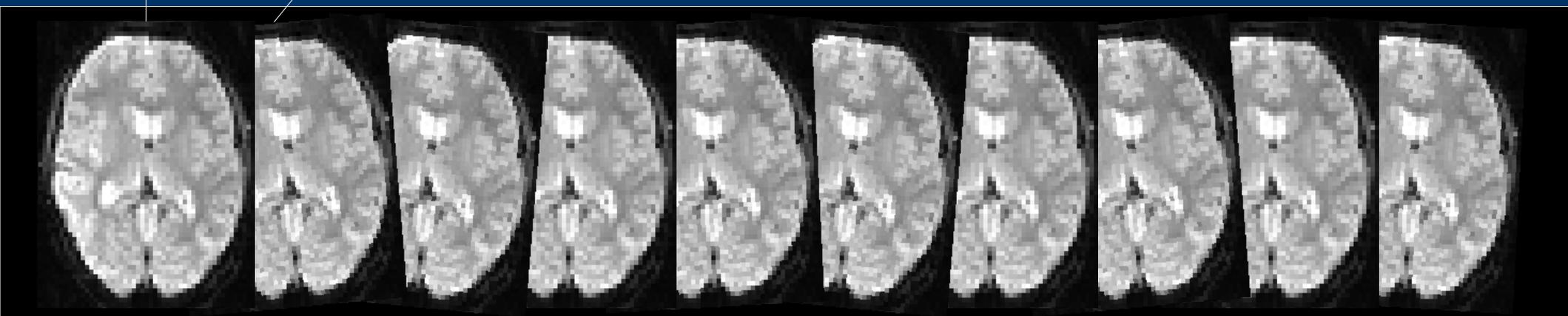
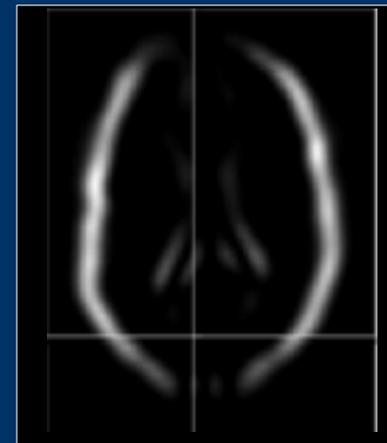
=

Difference



$^2 =$

Variance



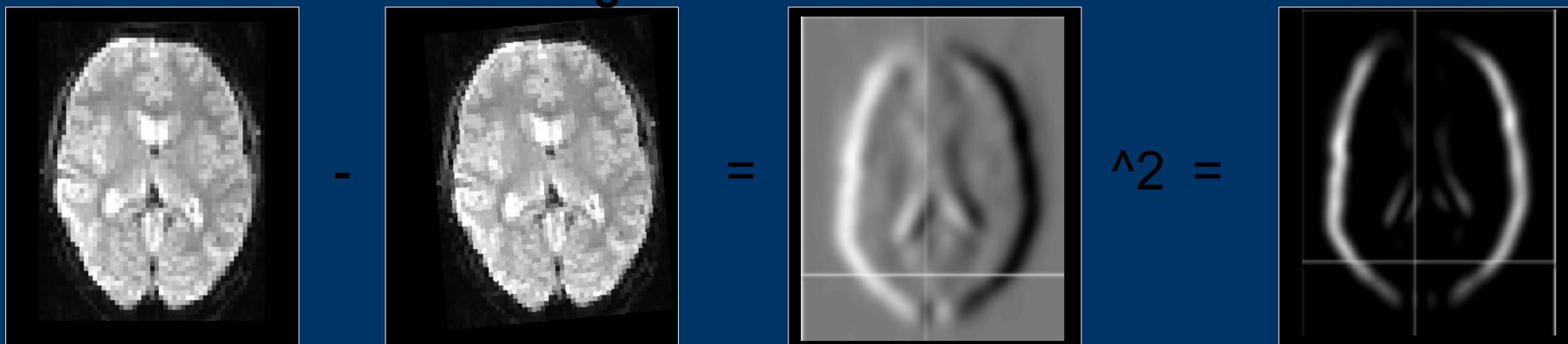
Motion Correction

Reference

I th Image

Difference

Variance



Rigid body (6dof)

Rigid body transformations parameterised by:

Translations

$$\begin{pmatrix} 1 & 0 & 0 & X_{\text{trans}} \\ 0 & 1 & 0 & Y_{\text{trans}} \\ 0 & 0 & 1 & Z_{\text{trans}} \\ 0 & 0 & 0 & 1 \end{pmatrix}$$

Pitch

$$\begin{pmatrix} 1 & 0 & 0 & 0 \\ 0 & \cos(\Phi) & \sin(\Phi) & 0 \\ 0 & -\sin(\Phi) & \cos(\Phi) & 0 \\ 0 & 0 & 0 & 1 \end{pmatrix}$$

Roll

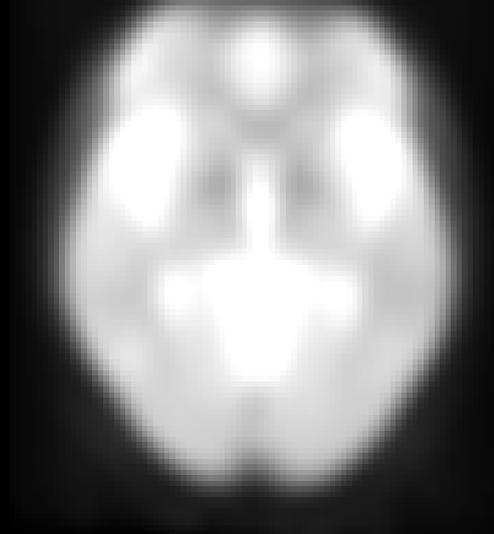
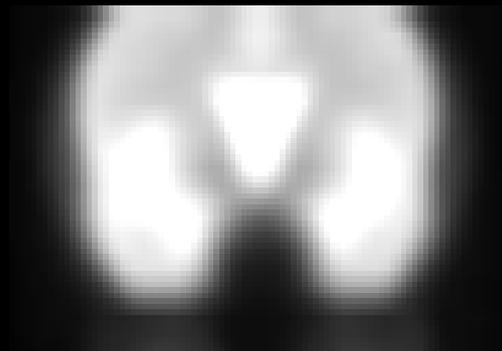
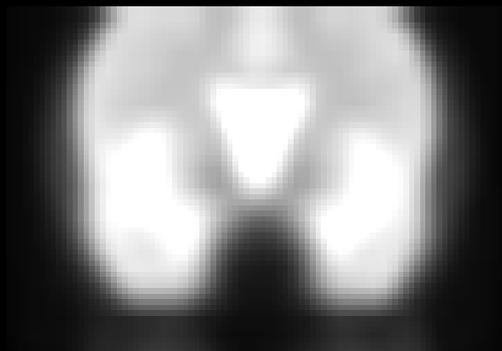
$$\begin{pmatrix} \cos(\Theta) & 0 & \sin(\Theta) & 0 \\ 0 & 1 & 0 & 0 \\ -\sin(\Theta) & 0 & \cos(\Theta) & 0 \\ 0 & 0 & 0 & 1 \end{pmatrix}$$

Yaw

$$\begin{pmatrix} \cos(\Omega) & \sin(\Omega) & 0 & 0 \\ -\sin(\Omega) & \cos(\Omega) & 0 & 0 \\ 0 & 0 & 1 & 0 \\ 0 & 0 & 0 & 1 \end{pmatrix}$$

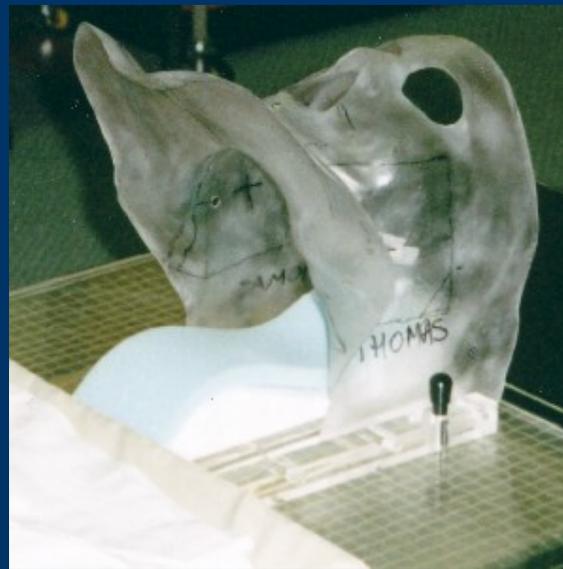
Viewing motion correction



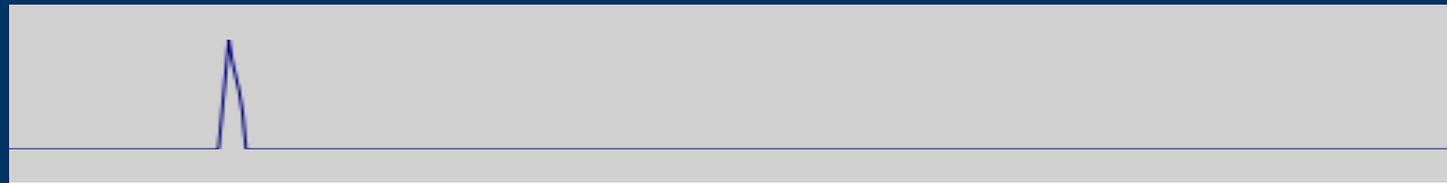
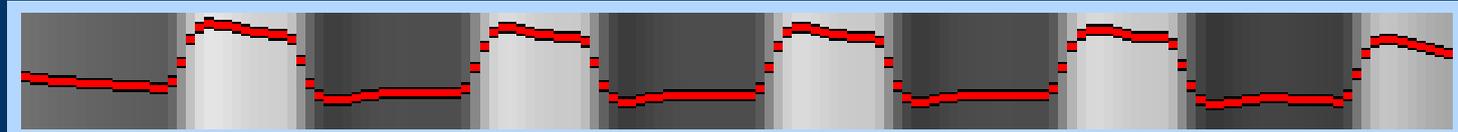
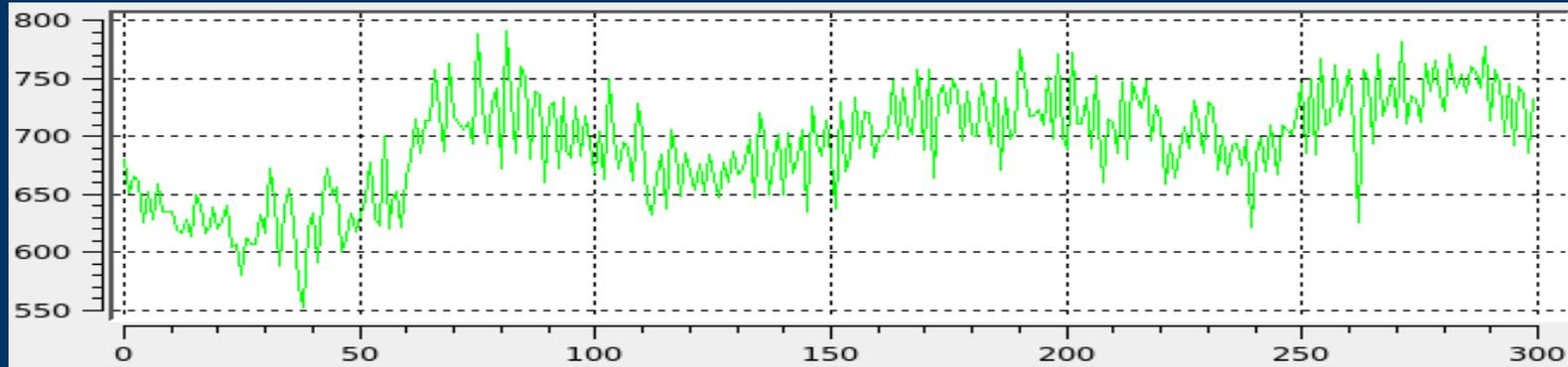
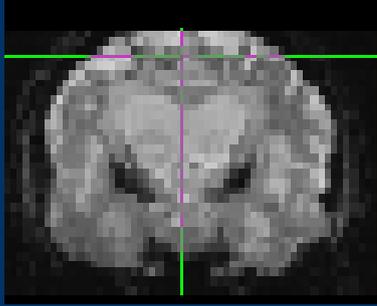


From FSL website

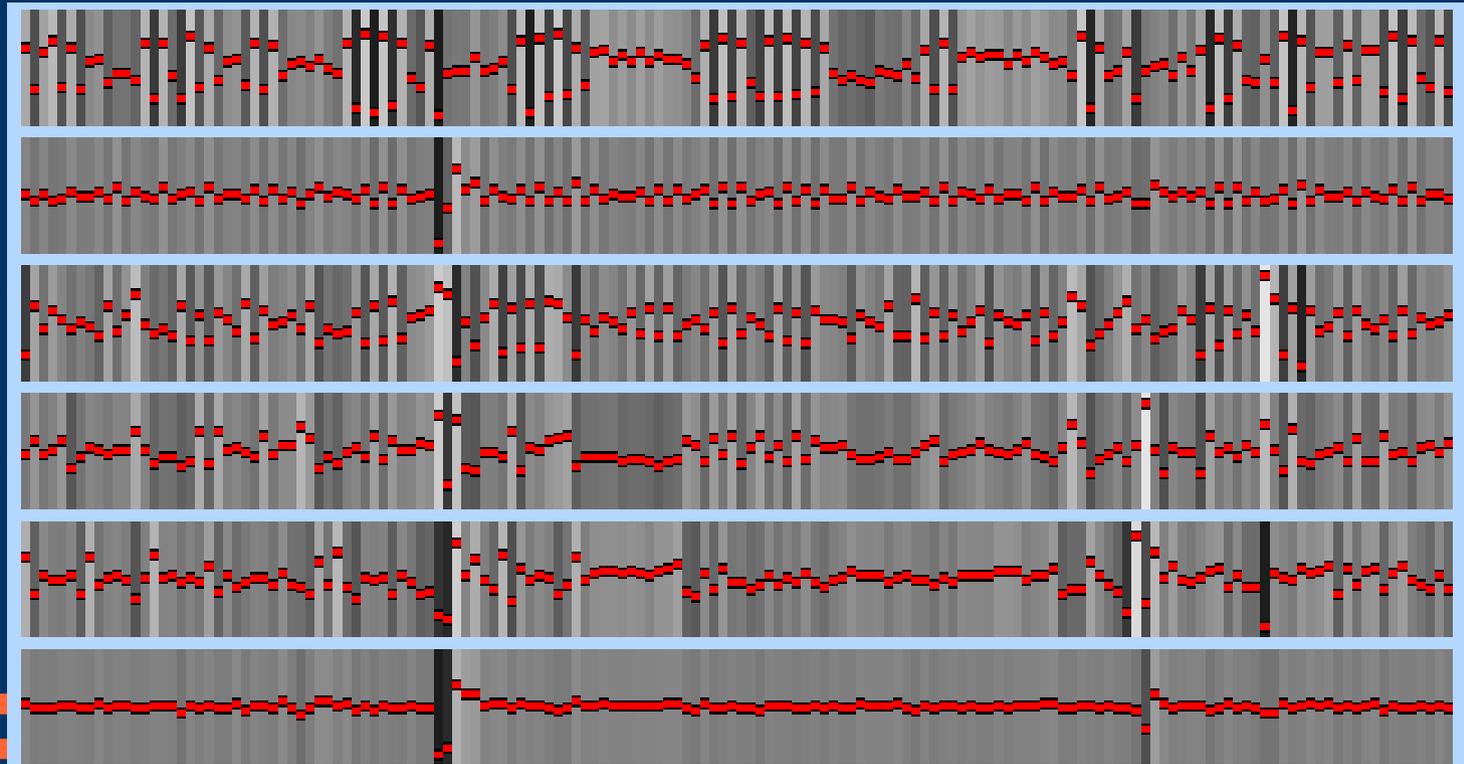
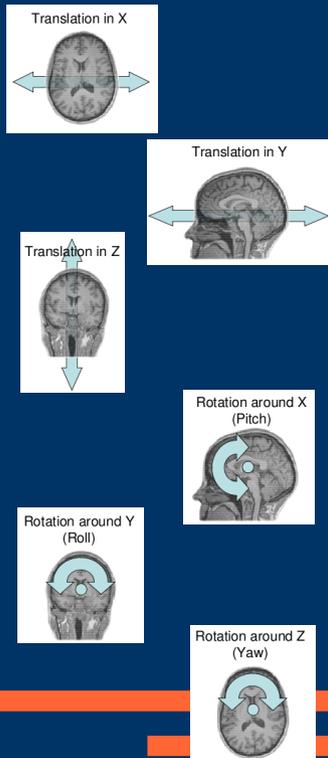
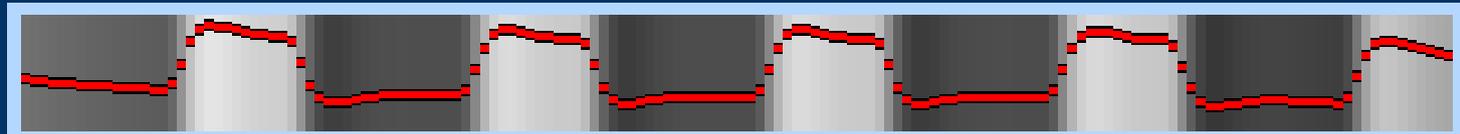
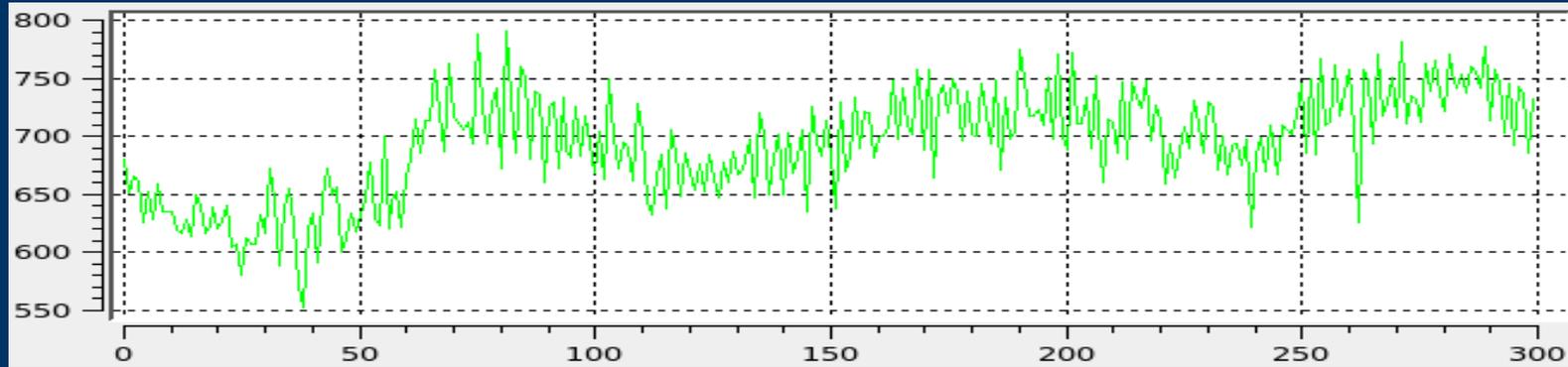
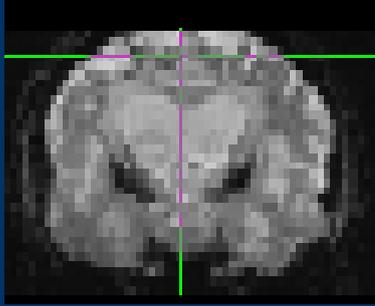
Coping with motion I: prevent it



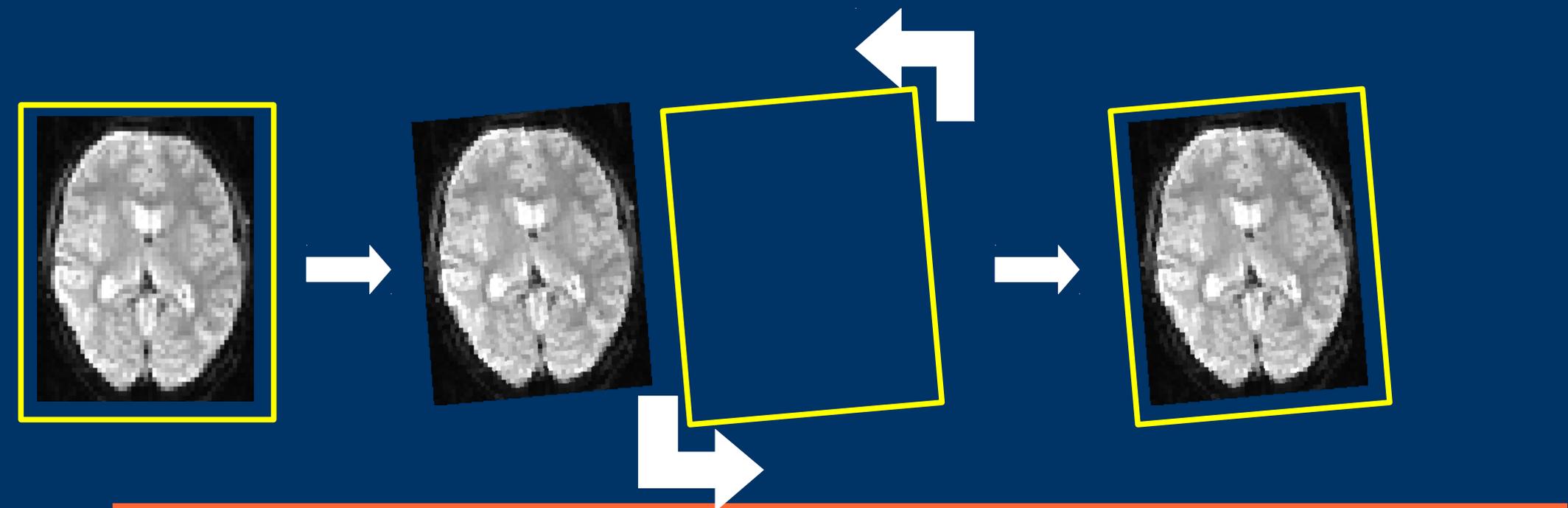
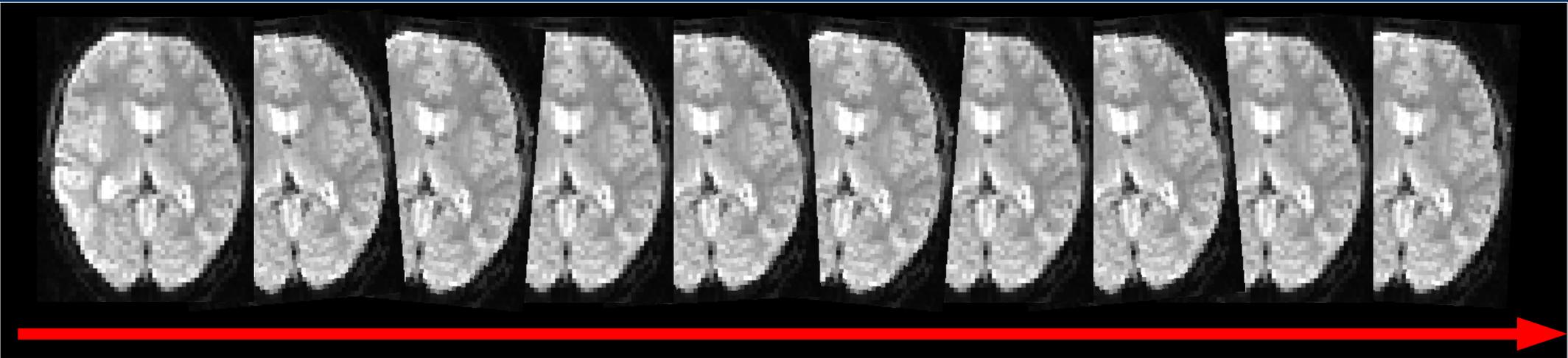
Coping with motion II(a): model it



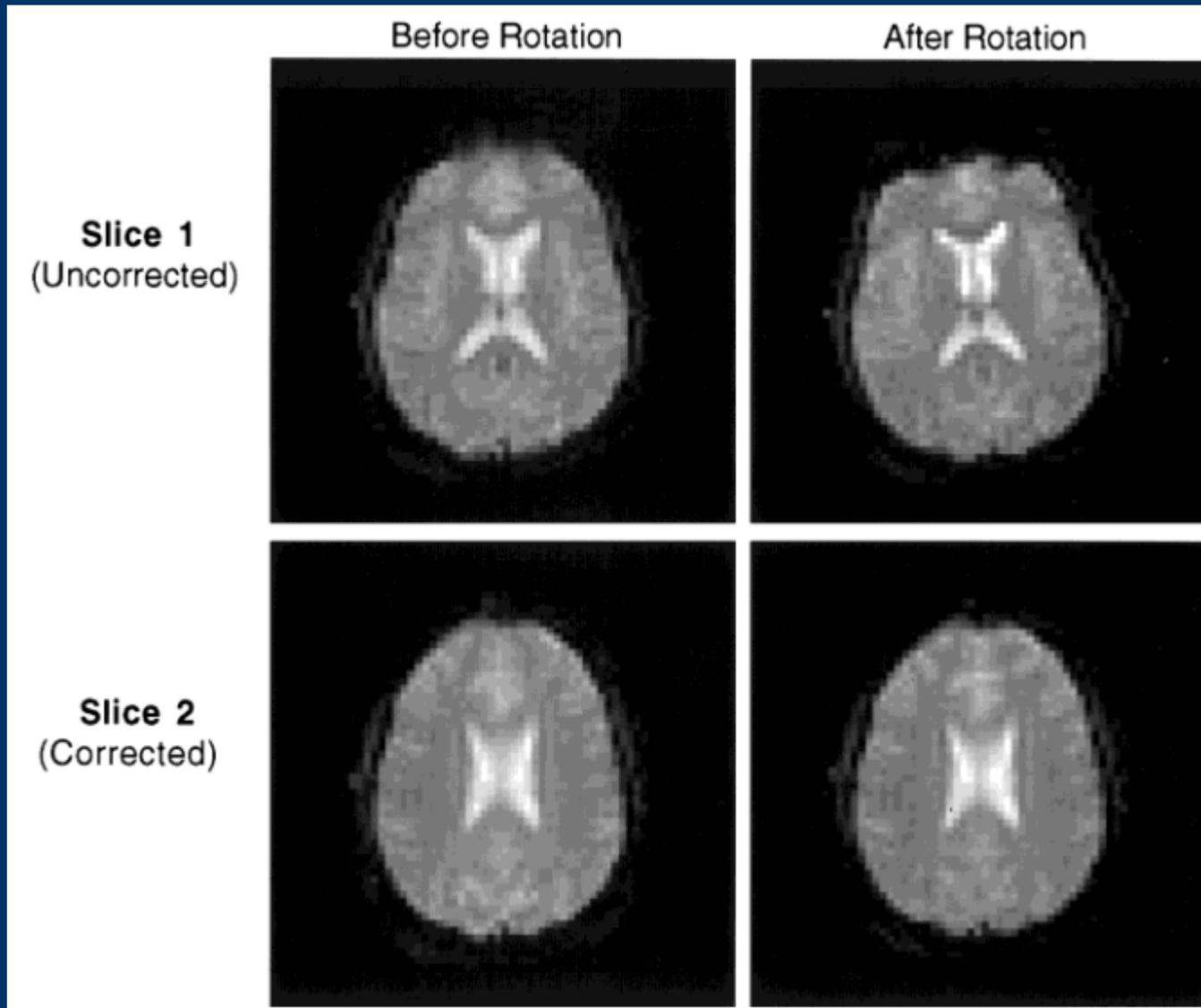
Coping with motion II(b): model it



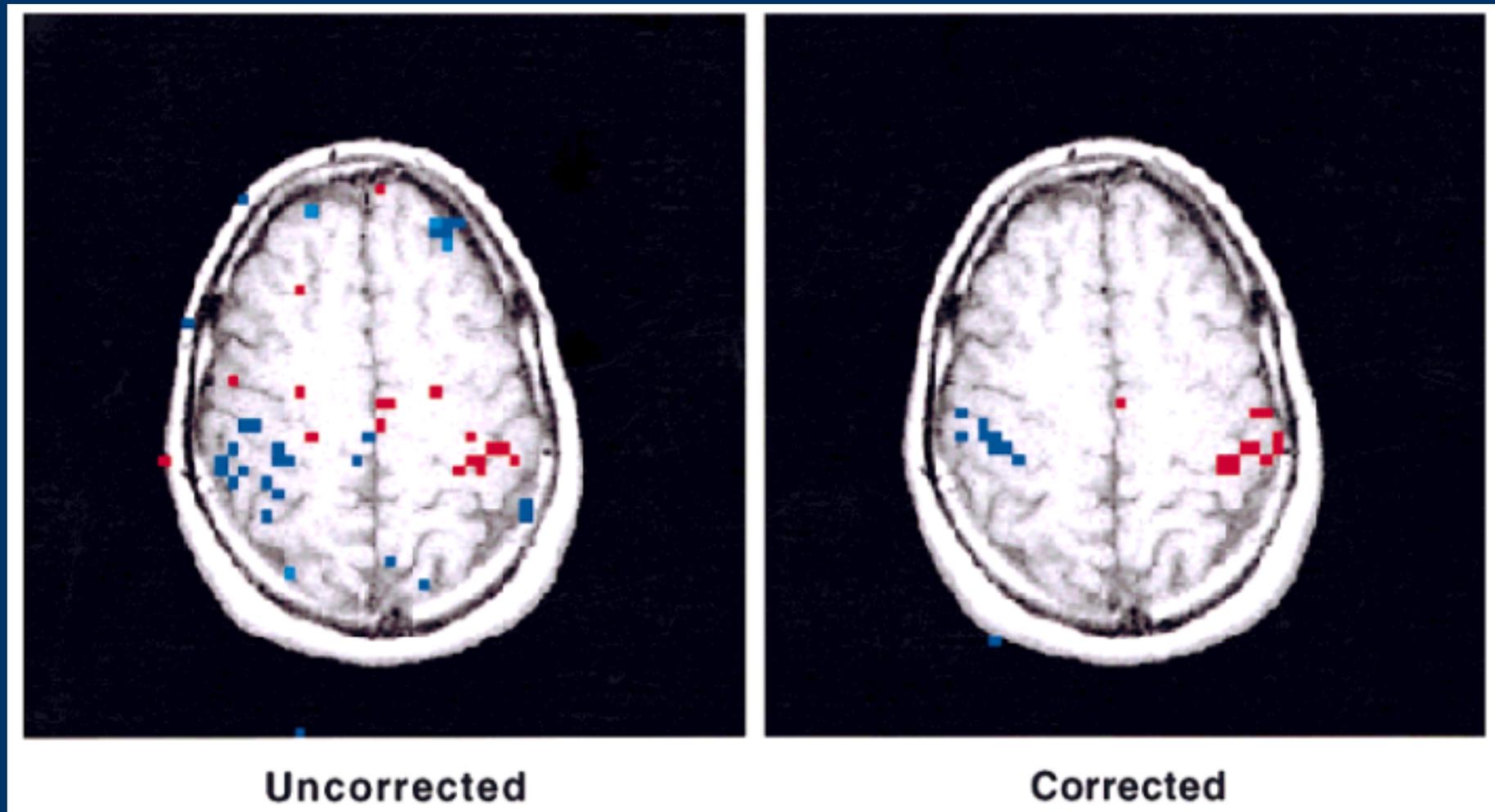
Coping with motion III: Prospective MC



Prospective motion correction



Prospective motion correction



Motion correction is good, however:

- Even after all this, movement artefacts still remain
 - Residual (uncorrected) motion
 - There's no way of detecting rapid movements within a scan
 - Spin history effects*
 - Voxels will be in different magnetic fields as a result*
 - Task correlated motion



The moral of the story...

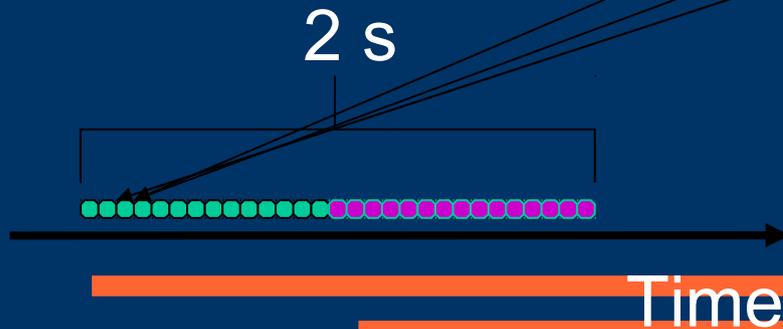
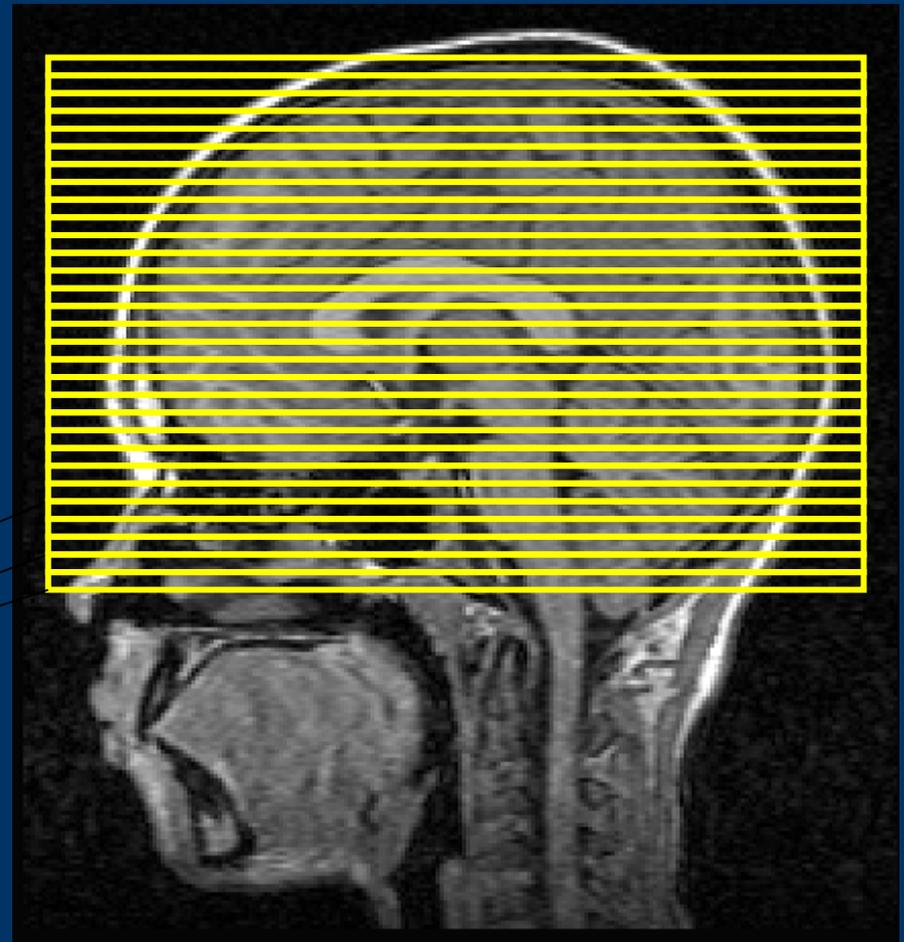
- *Stop people from moving*
 - Make sure they're comfortable to begin with
 - Tell them that motion is a big problem
 - Train subjects?
 - Reward them?
 - Decouple motion-prone tasks from cognitive event of interest
 - Model motion out
 - Reject run/subject
-
-

Preprocessing

- i. Motion correction
 - ii. Slice timing correction
 - iii. Spatial filtering
 - iv. Temporal filtering
 - v. Intensity normalization
 - vi. Field distortion correction (unwarping)
-
-

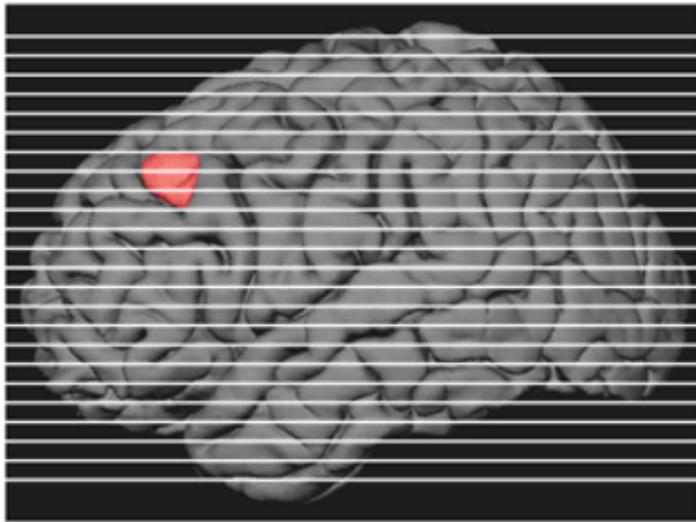
Slice timing correction

- In our exp we took a full functional image (volume) of the brain every 2 s.
- Each volume was acquired in 30 axial slices (interleaved).

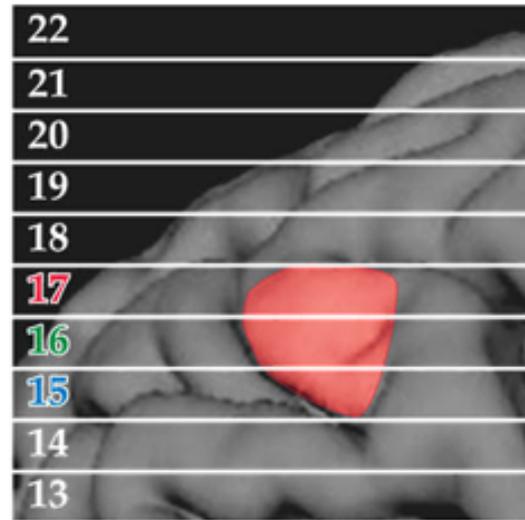


Slice timing correction

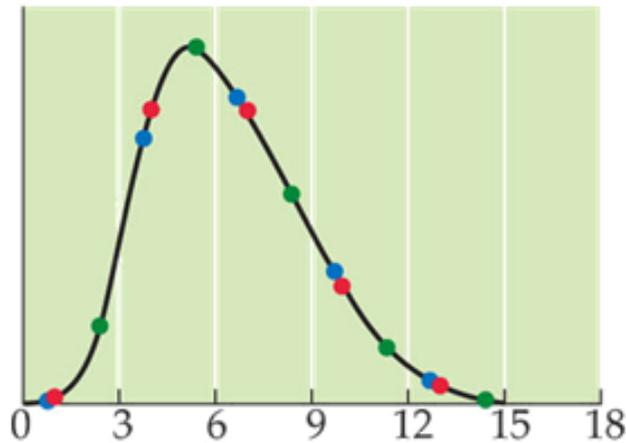
(A)



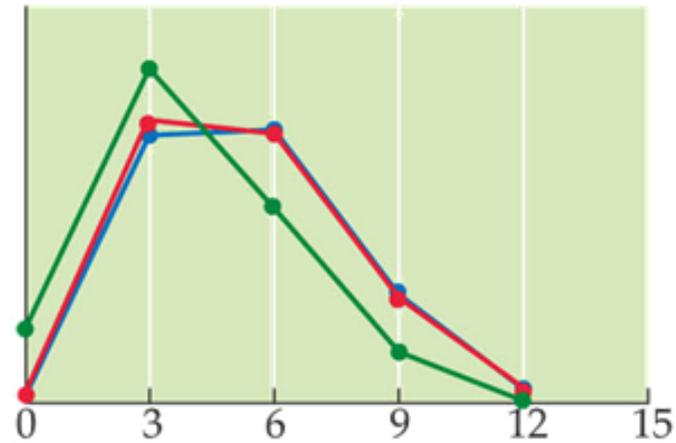
(B)



(C)

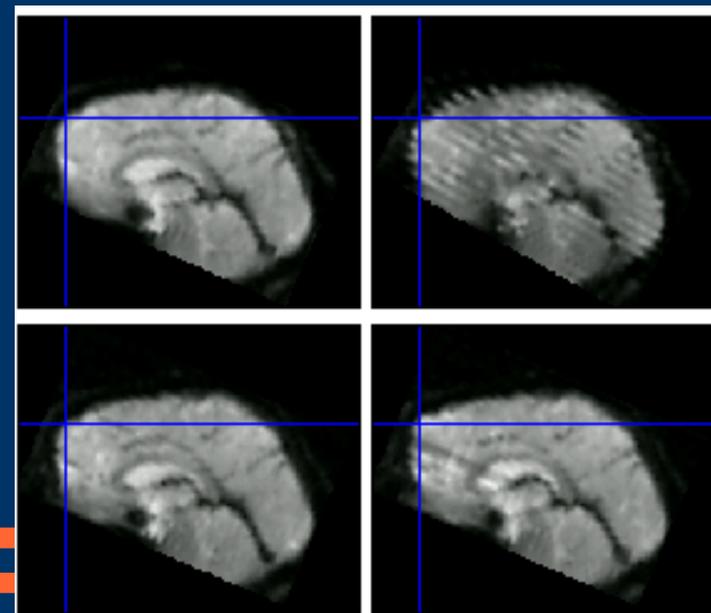


(D)



Slice timing correction

- Most people now suggest not to do it
 - i. Not all that helpful & requires interpolation
 - ii. It may worsen artefacts (e.g., smearing spikes)
 - iii. Interacts in unpredictable ways with motion correction
 - iv. We spatially smooth across proximal slices
 - v. Mismatching TR and task
 - vi. Include temporal derivative of HRF
- What order? Ascending, descending, contiguous, interleaved.



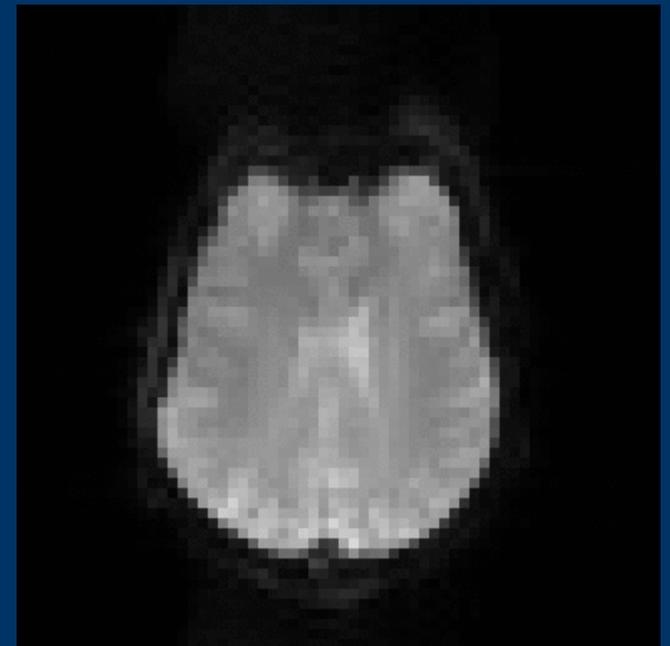
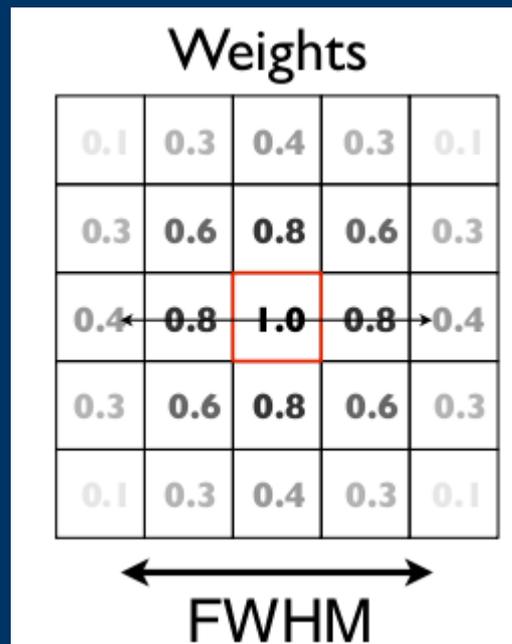
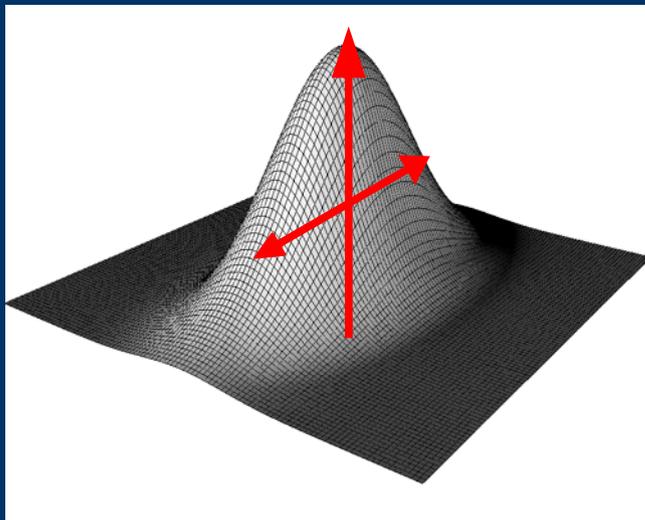
Preprocessing

- i. Motion correction
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- iv. Temporal filtering
- v. Intensity normalization
- vi. Field distortion correction (unwarping)



Spatial Filtering

- Replace each voxel's value with a weighted average of its value and the value of its neighbouring voxels.
- Gaussian kernel (mm FWHM)



Spatial Filtering

- Advantages

- Increases Signal to Noise Ratio (SNR)

- *Matched Filter Theorem*: Maximum increase in SNR by filter with same shape/size as signal

- Allows application of Gaussian Field Theory

- May improve comparisons across subjects

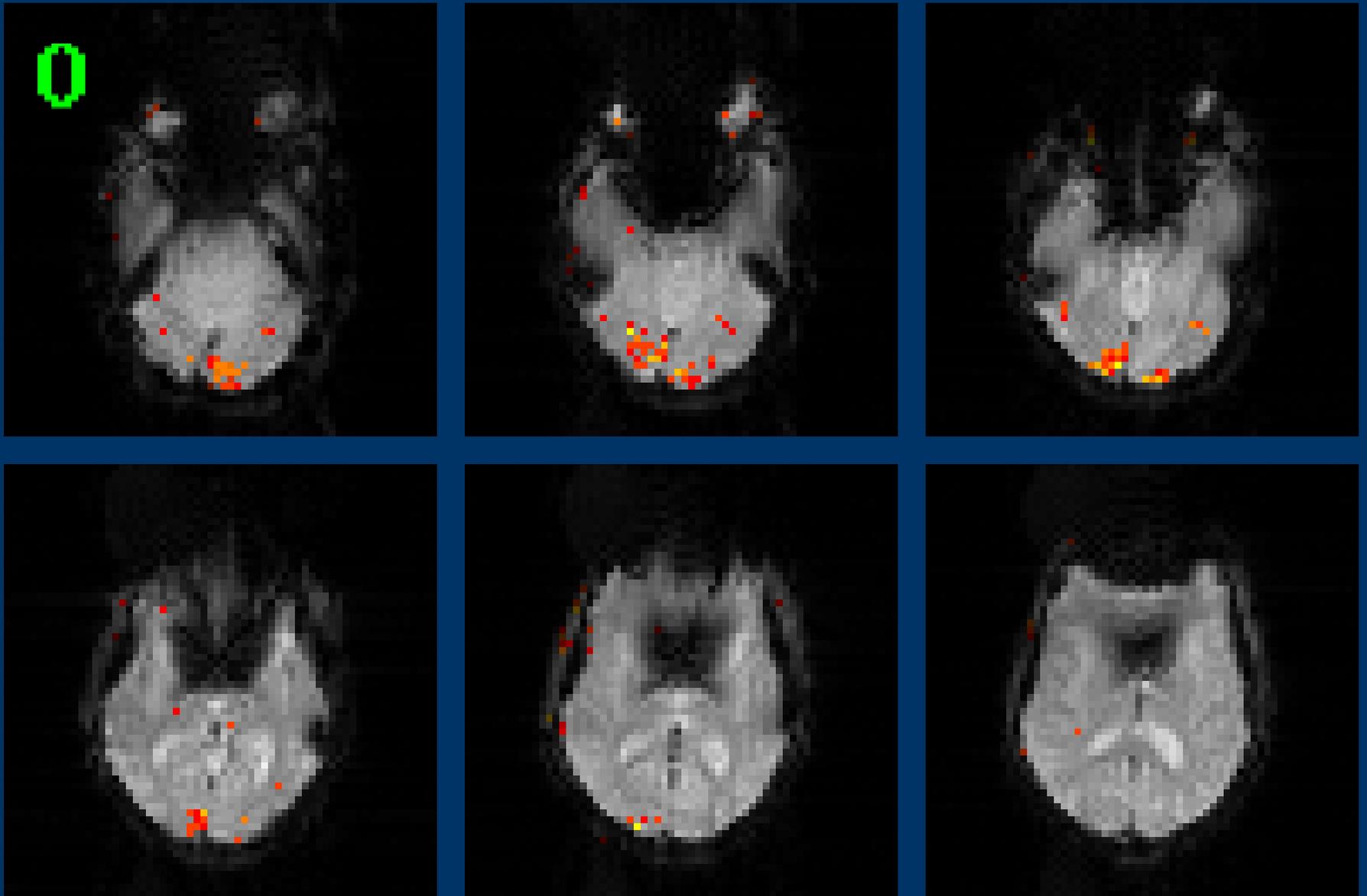
- Disadvantages

- Reduces spatial resolution

- May reduce your signal if smaller than your filter size!



Spatial Filtering



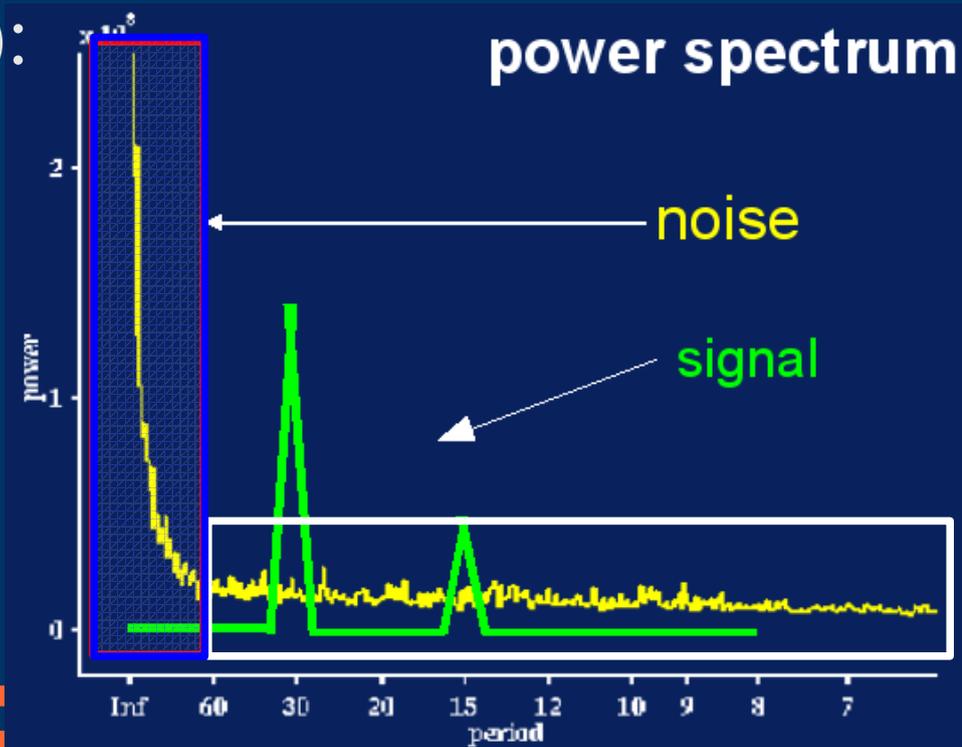
Preprocessing

- i. Motion correction
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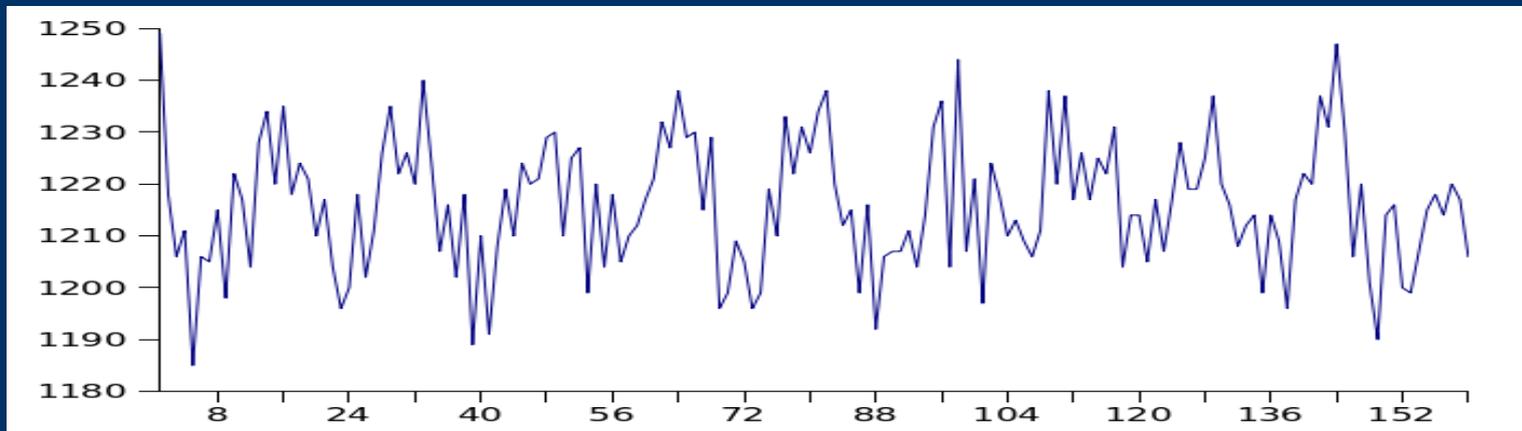
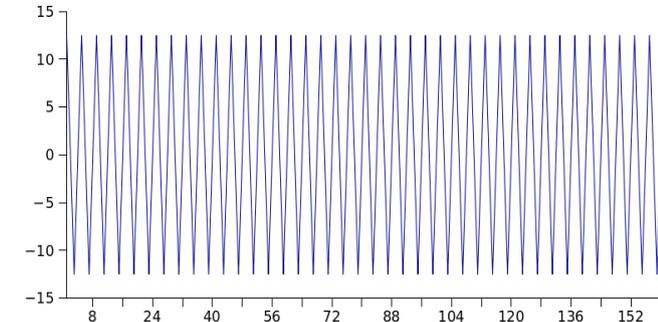
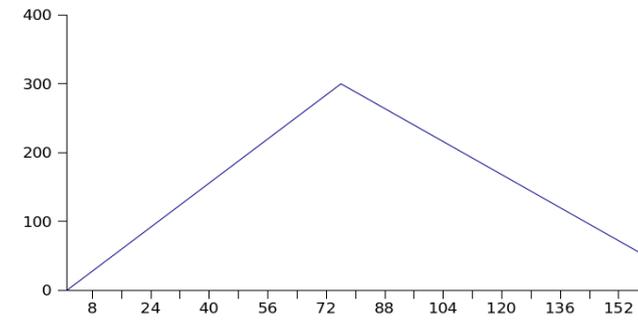
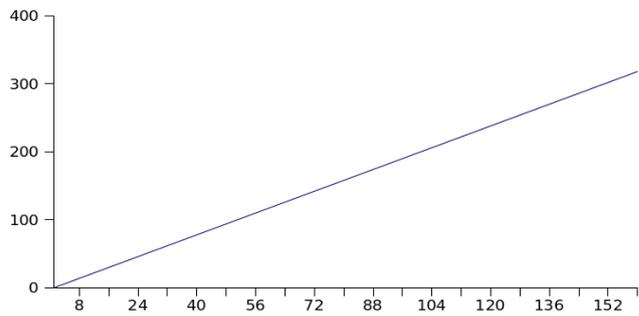
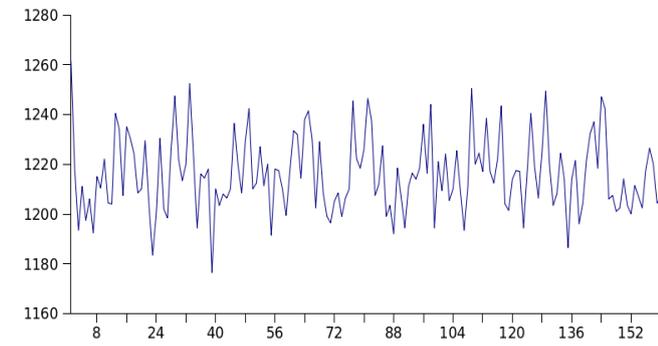
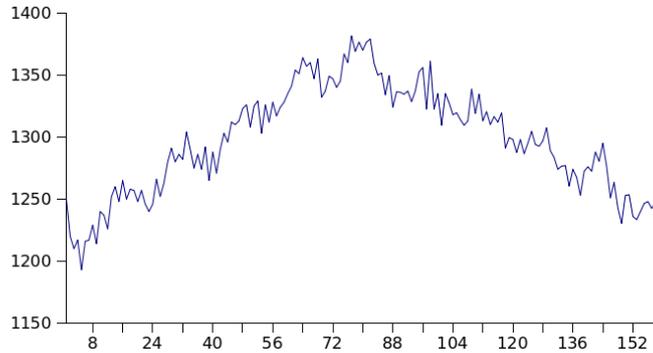
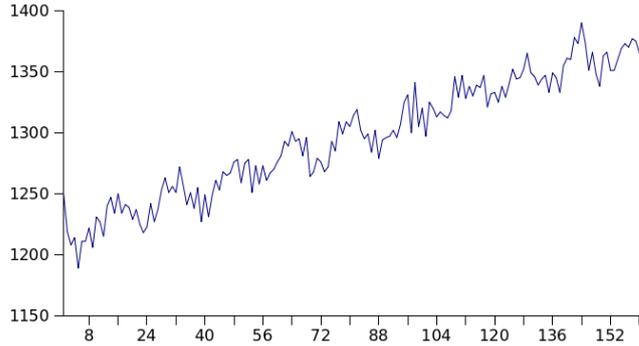


Temporal Filtering

- You are interested in the signal fluctuations that have to do with your task, and thus are at a specific frequency
- But there is a lot of activity at many other frequencies (particularly at low ones: $1/f$):
 - Scanner drift
 - Thermal noise
 - Heart beat
 - Respiration
 - Alertness
 - Learning

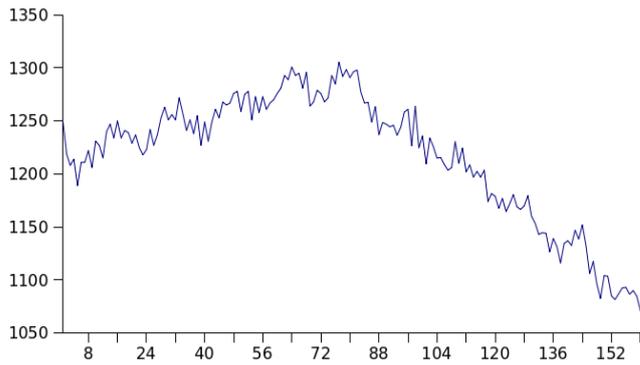


Signal & Noise

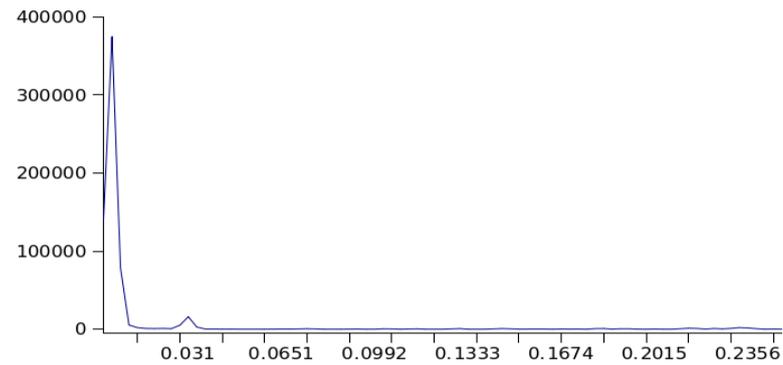


High-Pass Filtering

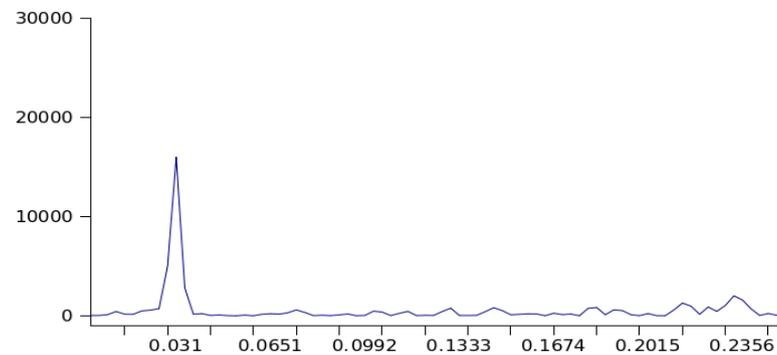
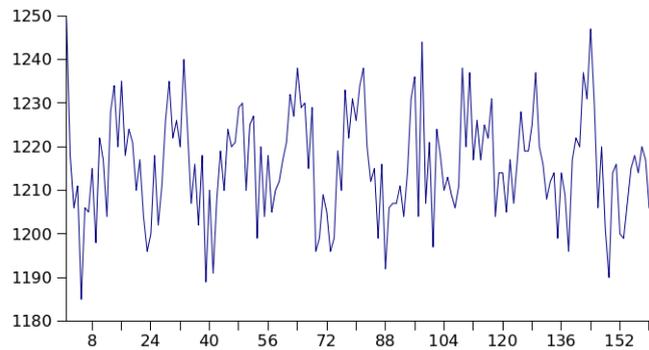
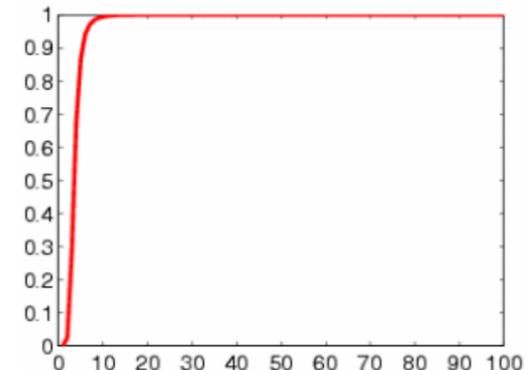
Timecourse



Power Spectrum

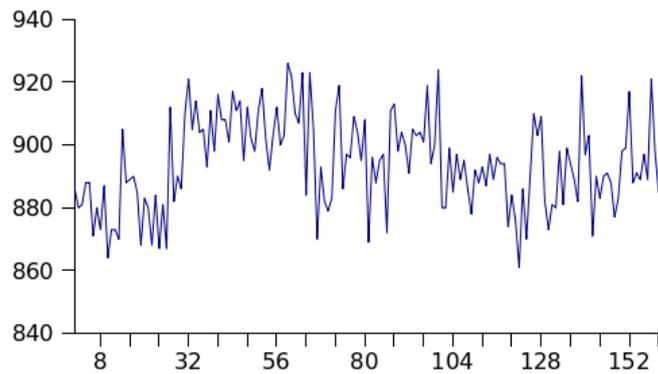


HP Filter

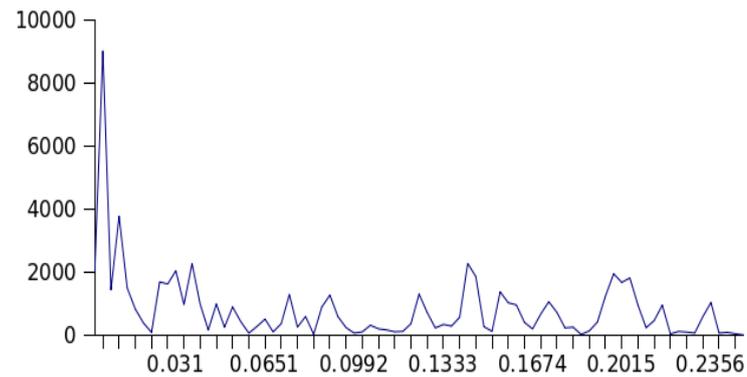


High-Pass Filtering

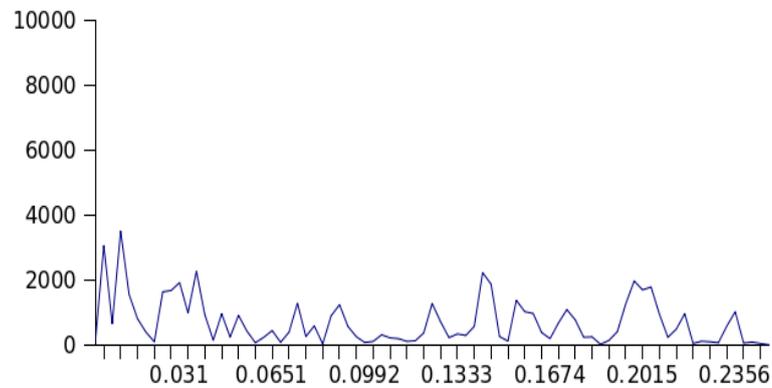
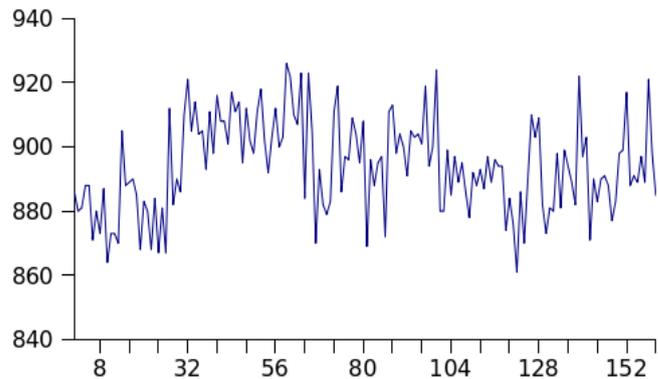
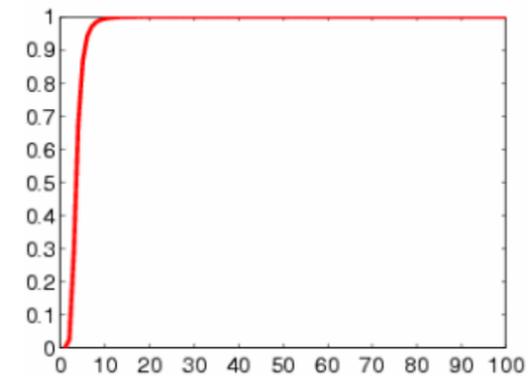
Timecourse



Power Spectrum

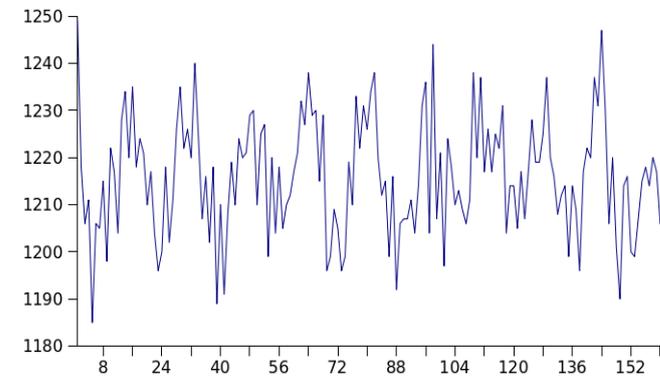
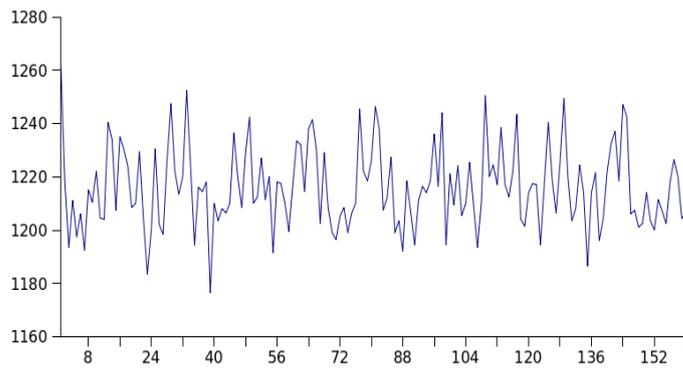


HP Filter

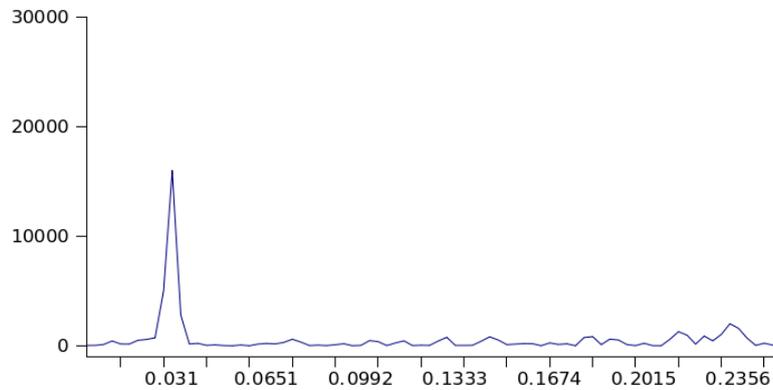
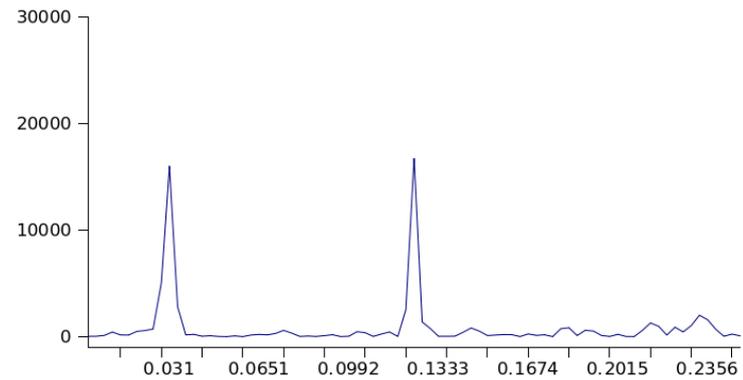


Low-Pass Filtering

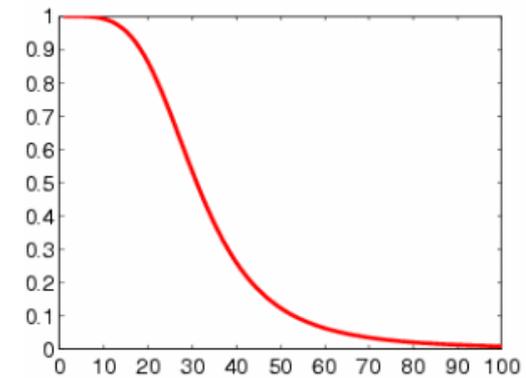
Timecourse



Power Spectrum

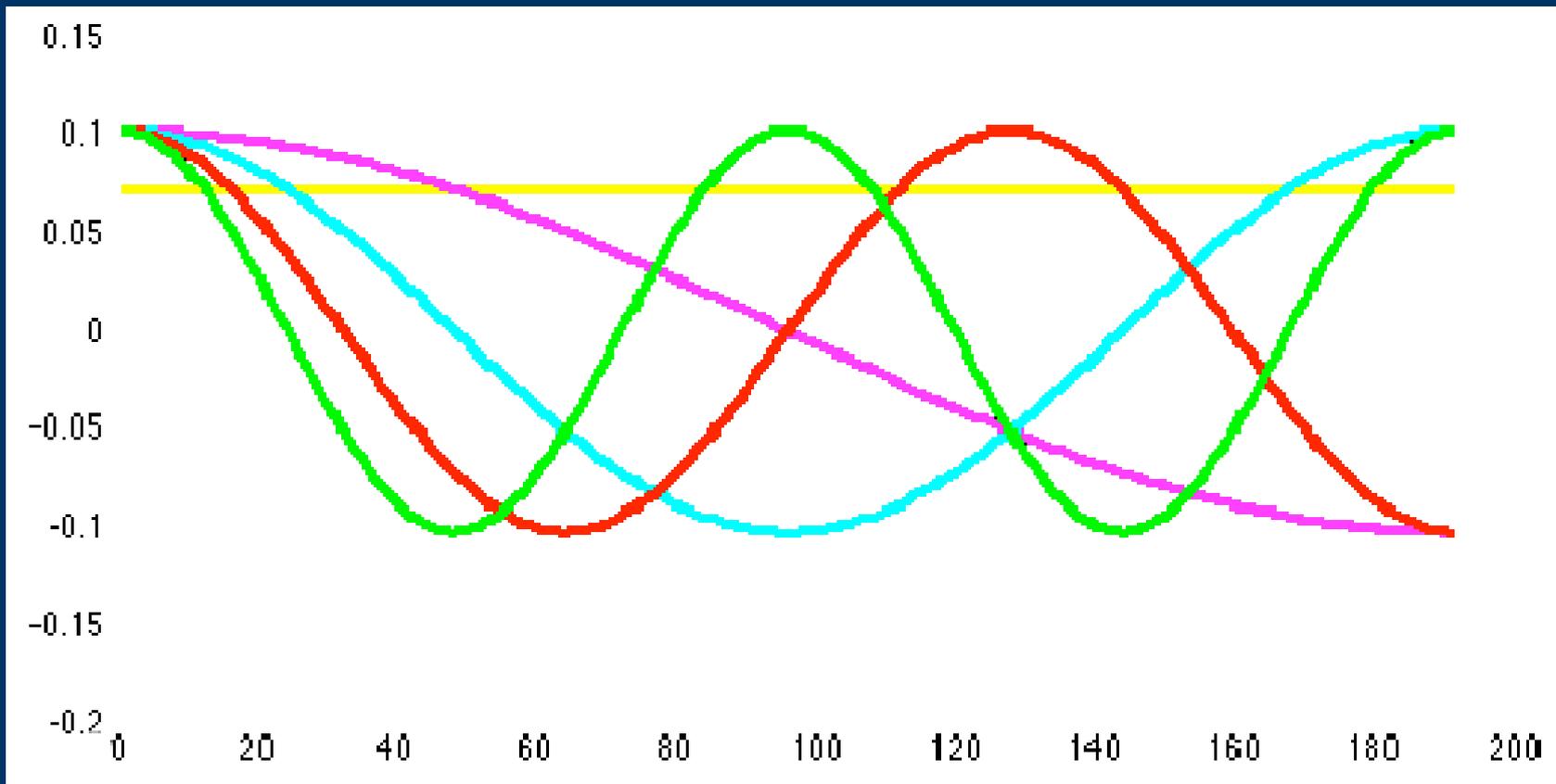


LP Filter



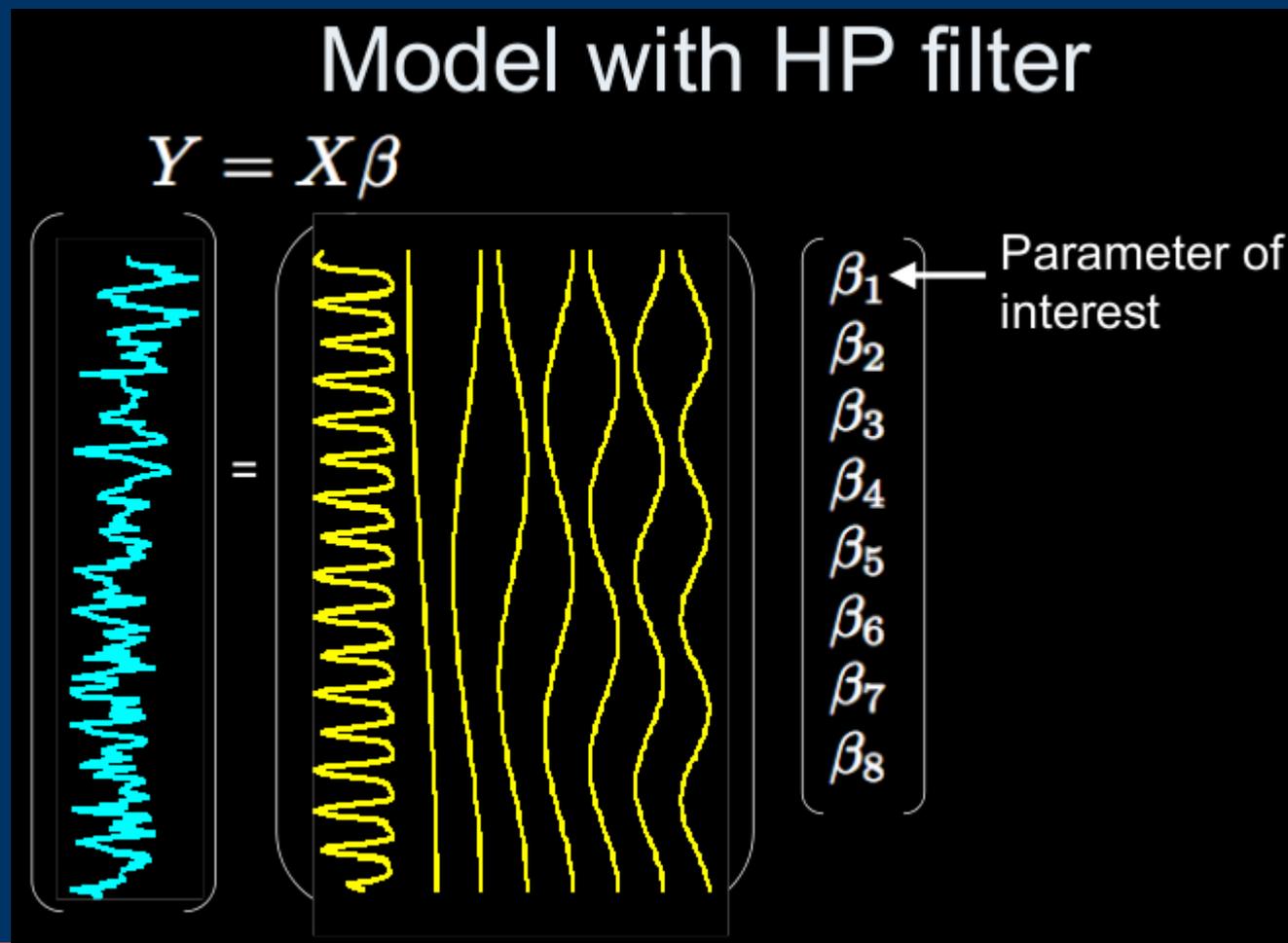
HP Filtering Strategy I: SPM

- Model low drifts to “soak up” their variance (using a discrete cosine transform basis set).



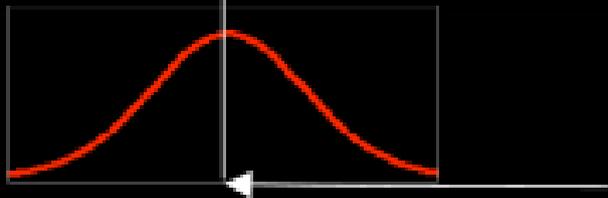
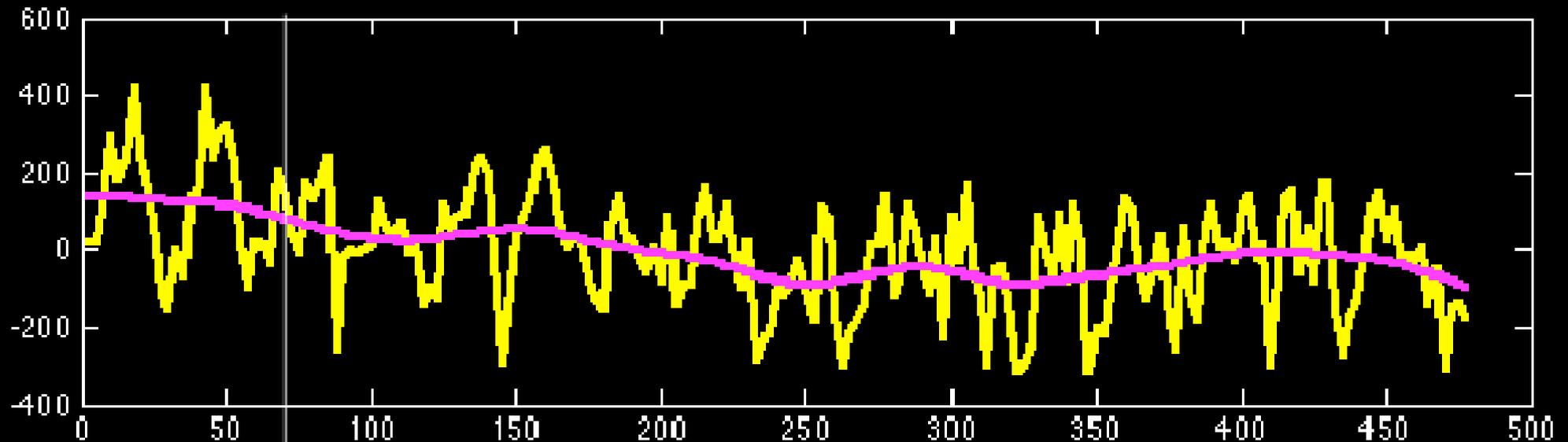
HP Filtering Strategy I: SPM

- Model low drifts to “soak up” their variance (using a discrete cosine transform basis set).



HP Filtering Strategy II: FSL

- Remove low drifts from the signal:
 - i. Fit a Gaussian-weighted running line



Fit at time t is a weighted average of data around t

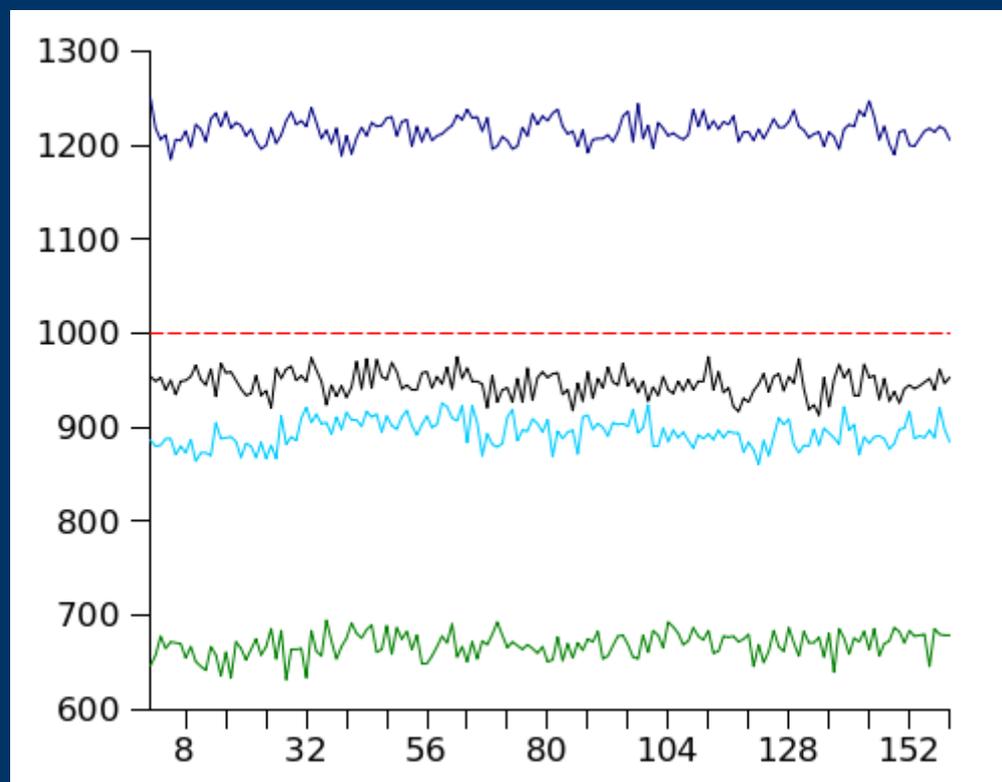
Preprocessing

- i. Motion correction
- ii. Slice timing correction
- iii. Spatial filtering
- iv. Temporal filtering
- v. Intensity normalization
- vi. Field distortion correction (unwarping)



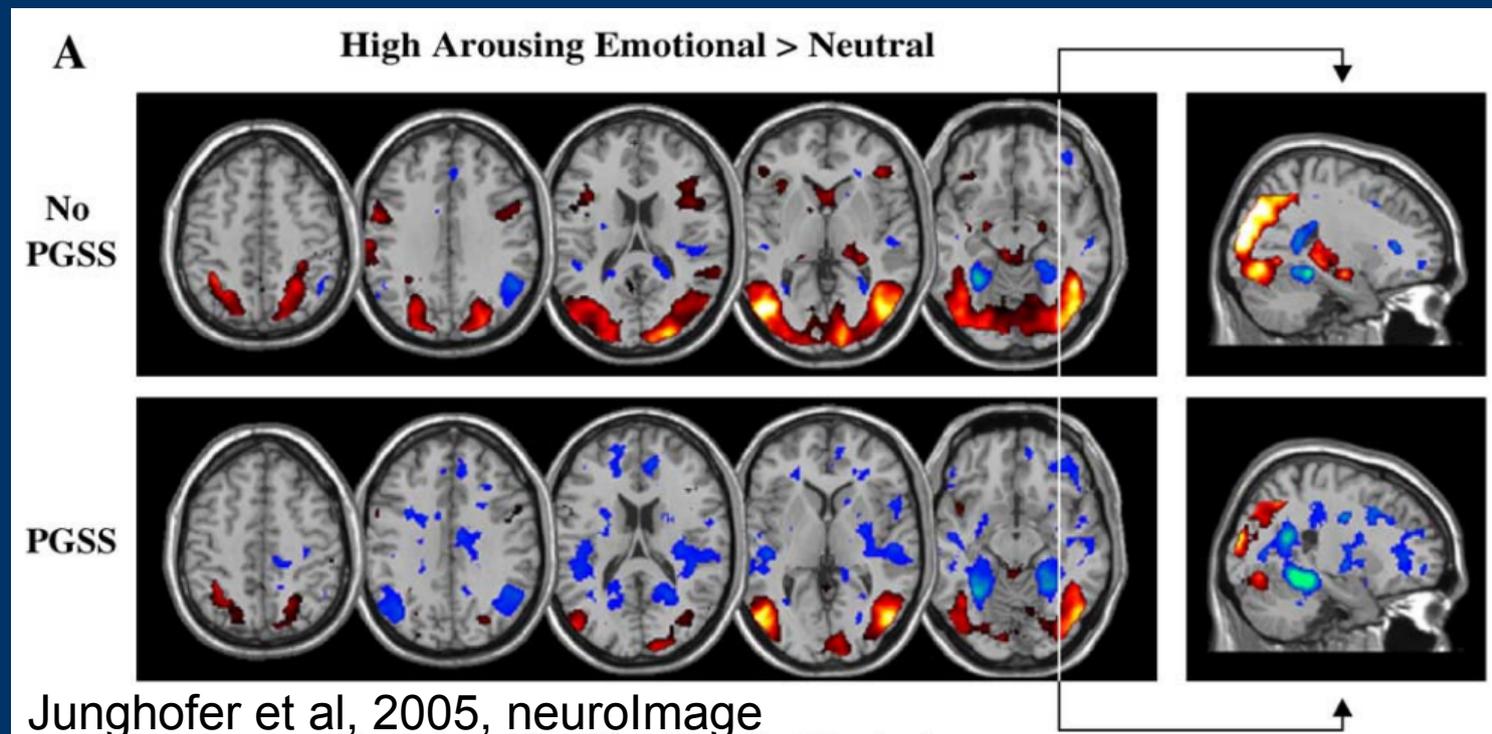
Intensity Normalization

- *Between-session (grand mean scaling)*
- The mean intensity of each 4D dataset varies for non-experimentally interesting reasons.
- Scale each 4D time-series by a single factor.
- Time-series from different runs are now centred around the same mean.



Intensity Normalization

- *Within-session*
- Forces each volumes (*within a run*) to have the same mean intensity.



Preprocessing

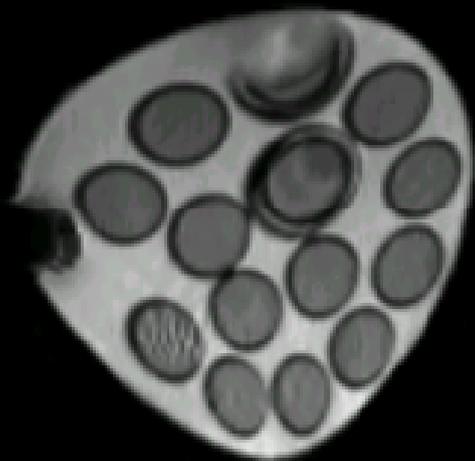
- i. Motion correction
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- iv. Temporal filtering
- v. Intensity normalization
- vi. Field distortion correction (unwarping)



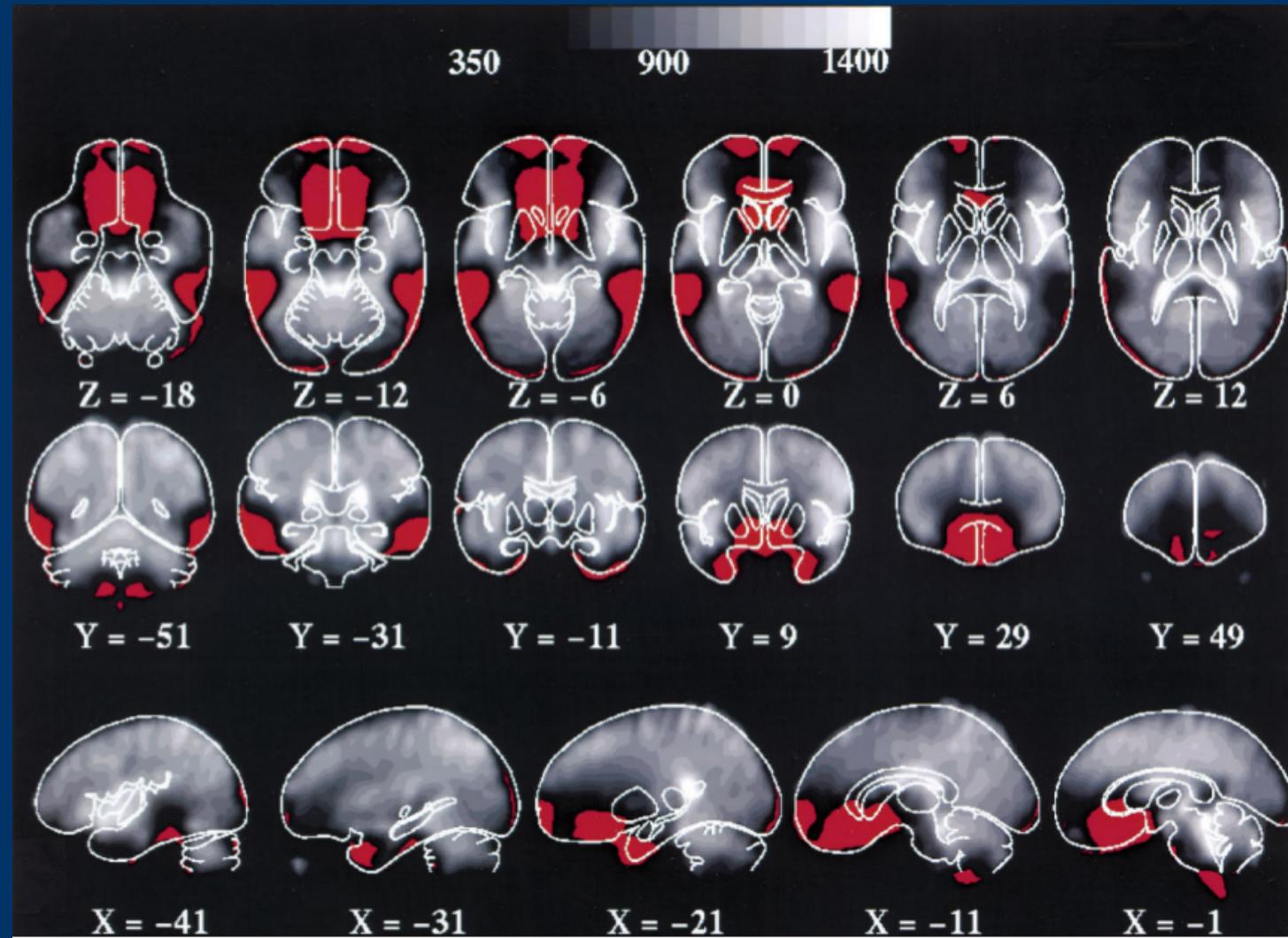
Geometric distortions



high res image

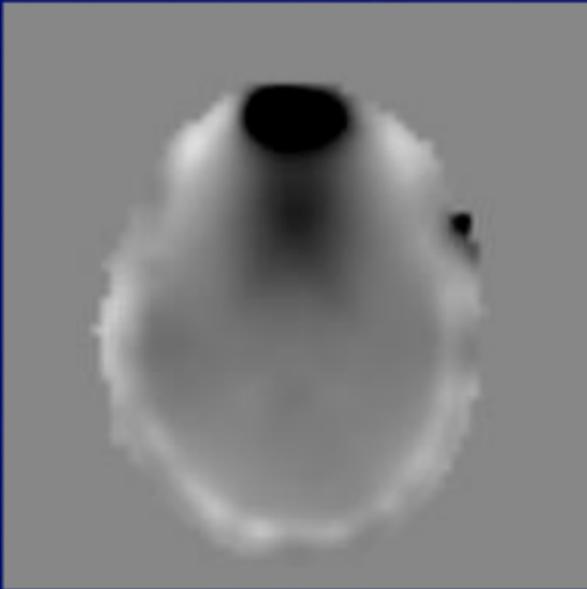


warped epi image

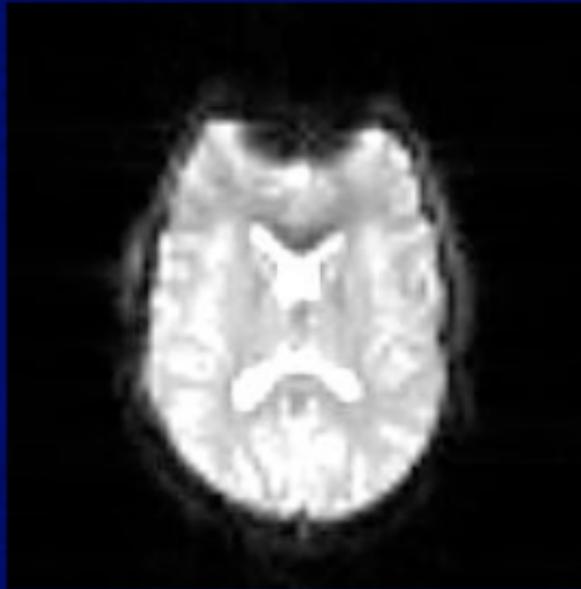


Un-warping for field distortion

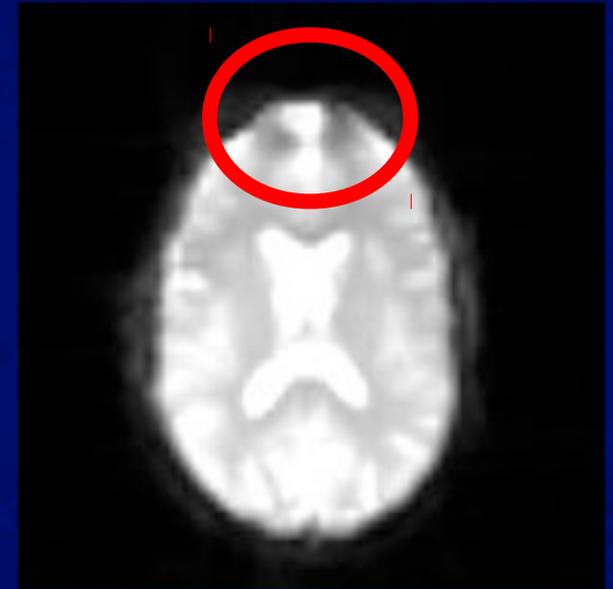
Field Map



EPI

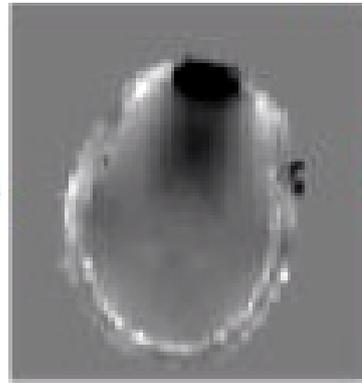


Unwarped EPI

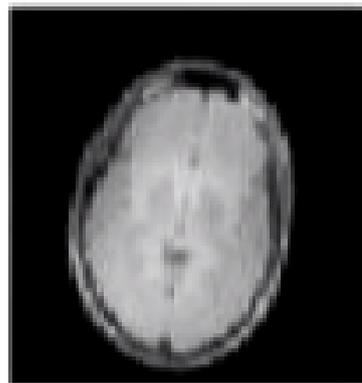


Fieldmap

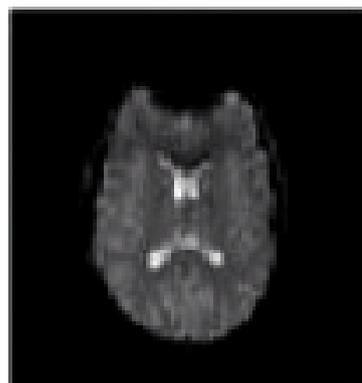
Phase



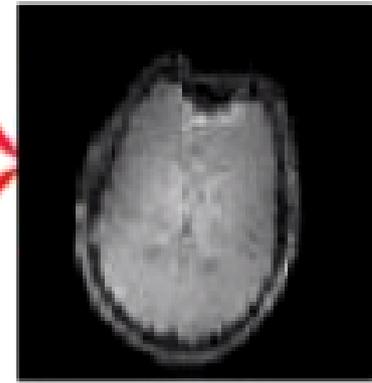
Abs



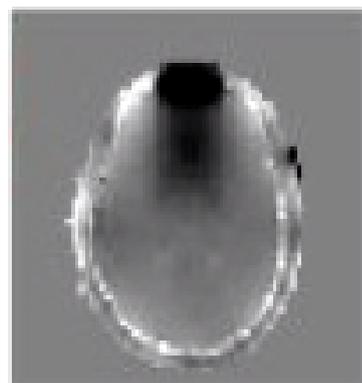
EPI



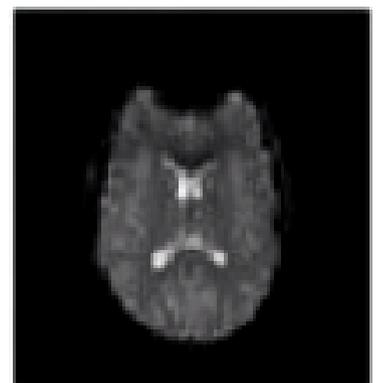
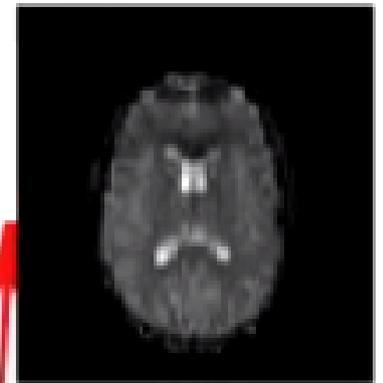
Forward Warp



Register 6 DOF



Unwarp

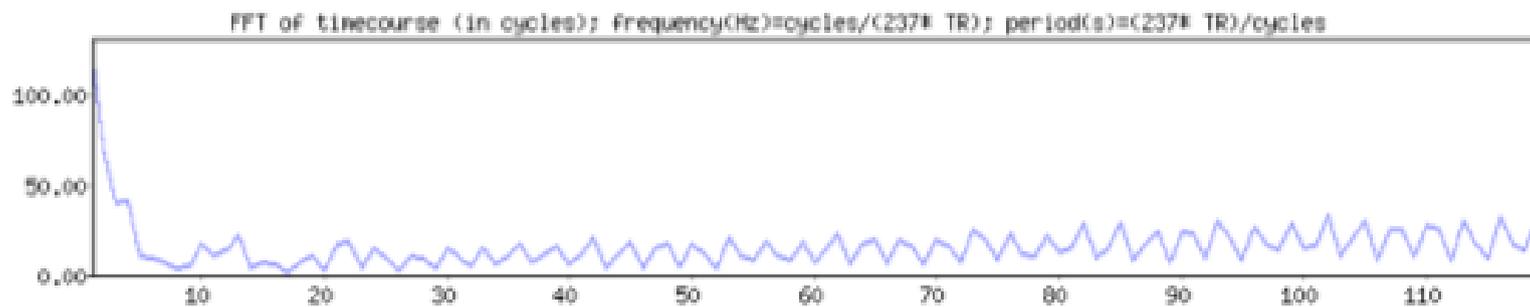
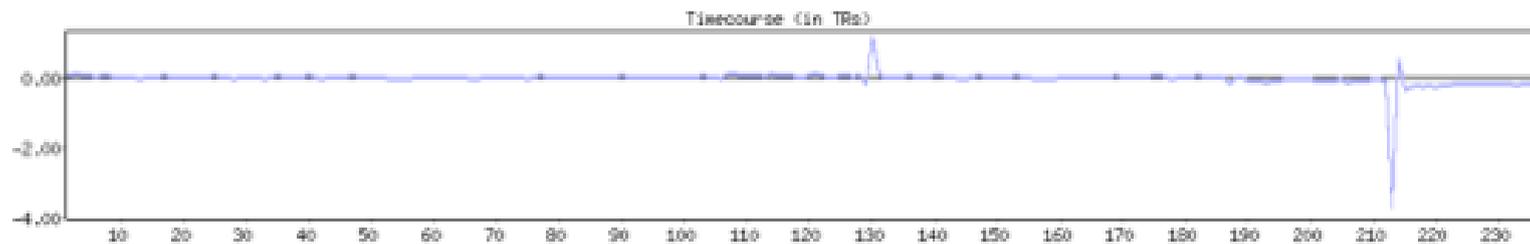
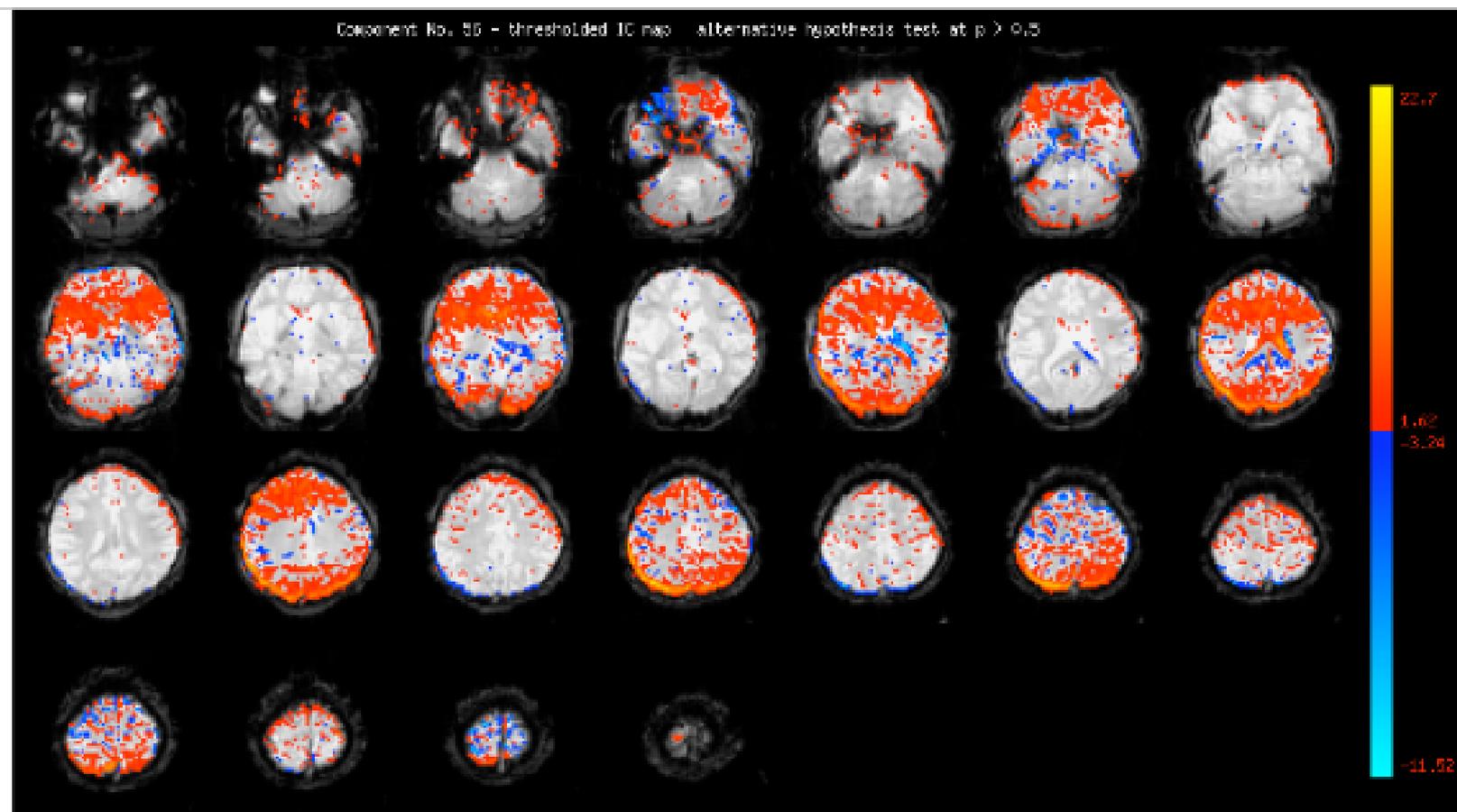


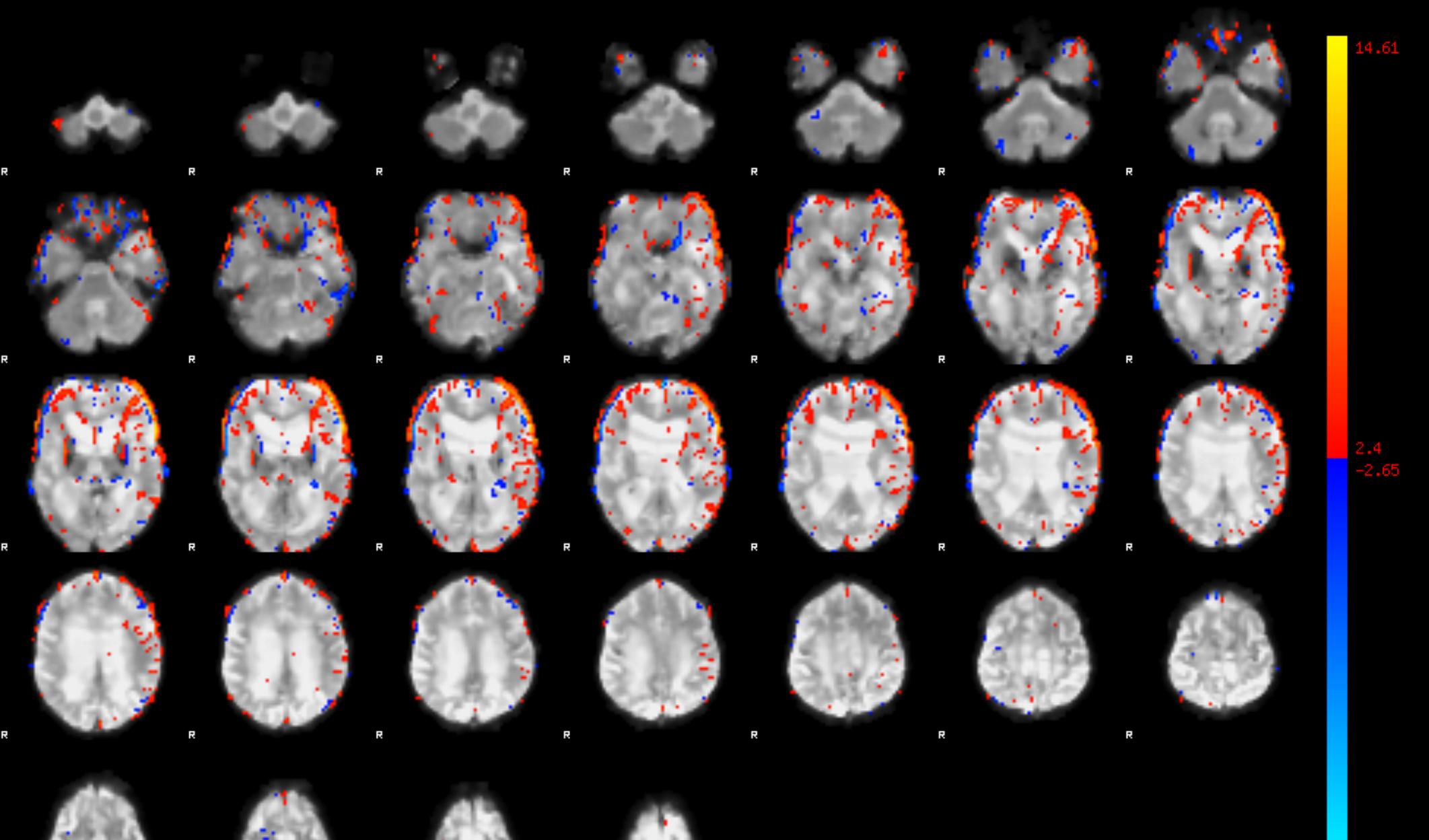
Preprocessing 2.0: ICA

1. Run an Independent Component Analysis
2. Find noise components
3. “Clean” the data
4. Run analysis on de-noised data

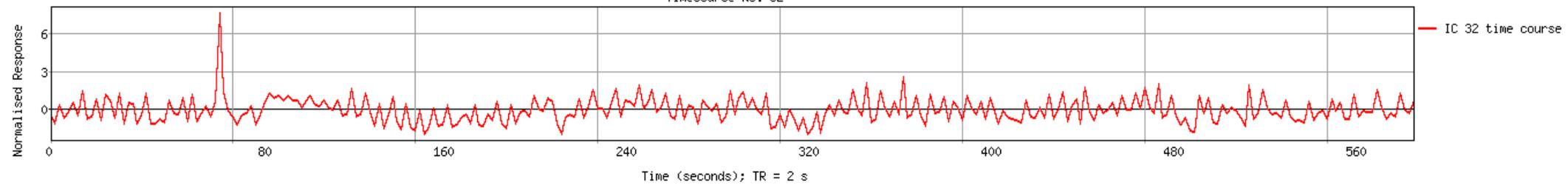


Abrupt motion



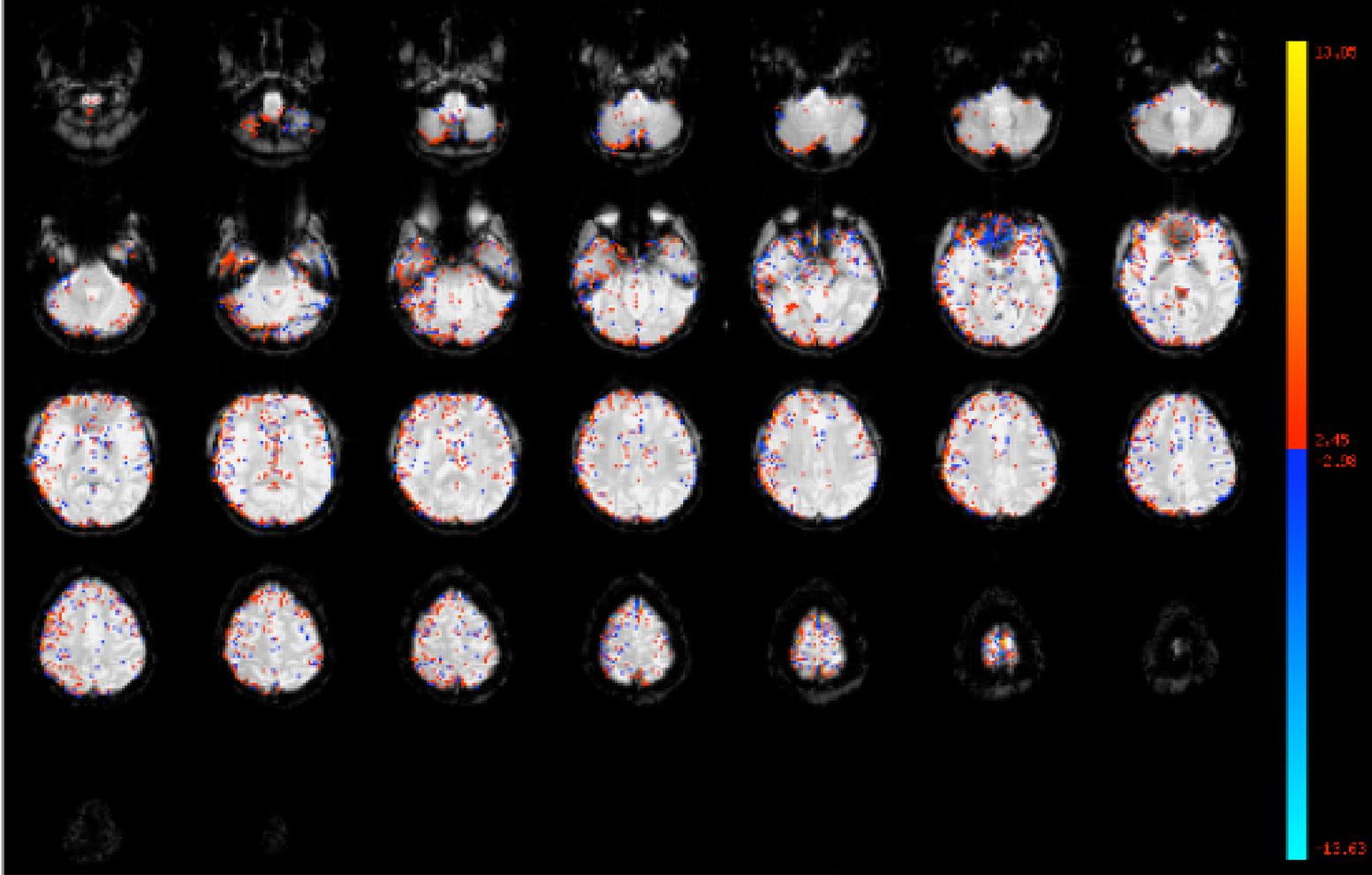


Timecourse No. 32

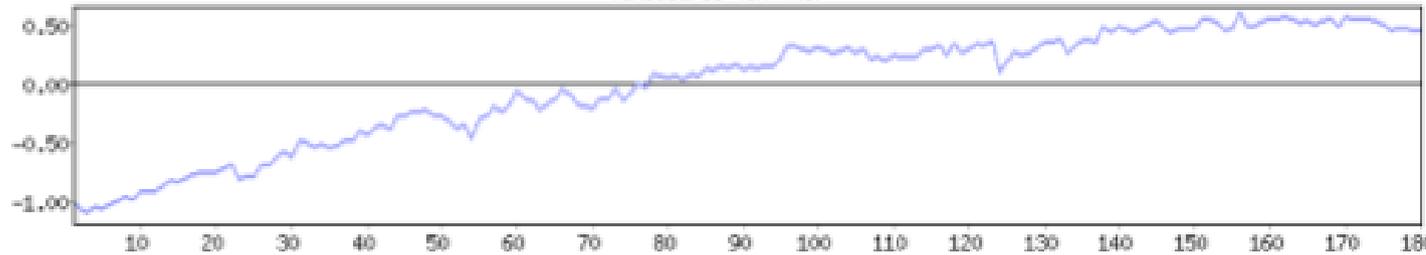


Slow drift

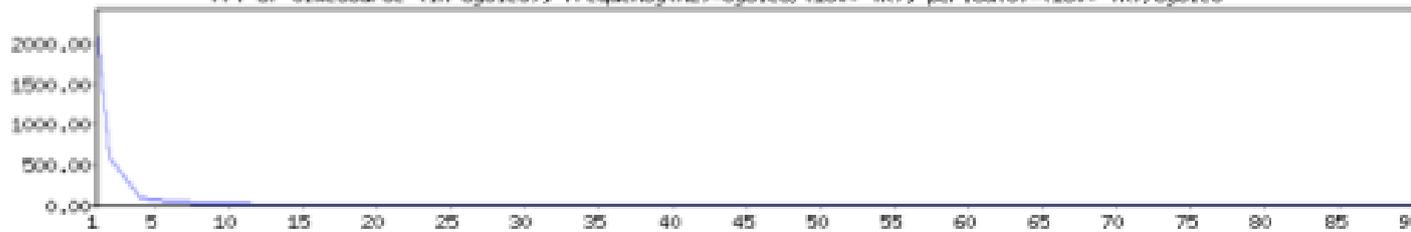
Component No. 22 - thresholded IC map - alternative hypothesis test at $p > 0.5$



Timecourse (in TRs)

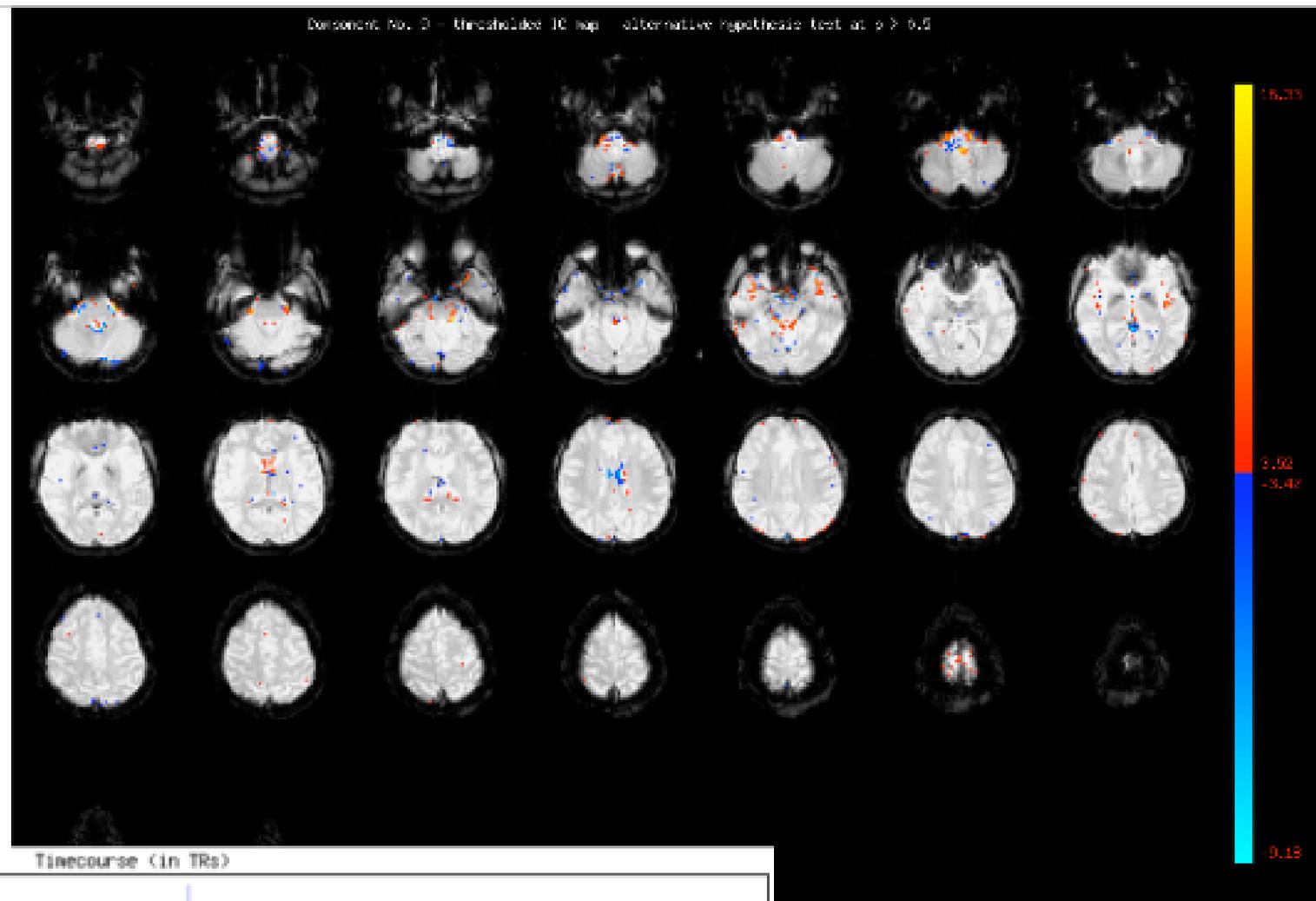


FFT of timecourse (in cycles): Frequency(Hz)=cycles/(180* TR); period(s)=(180* TR)/cycles

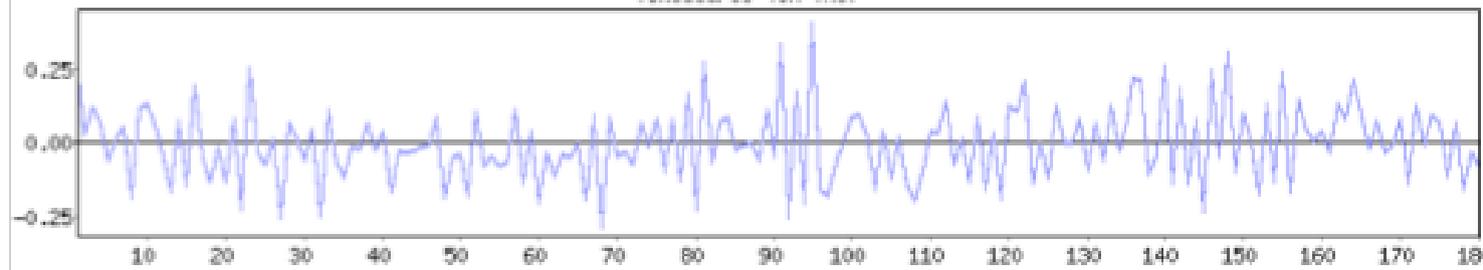


Source: R. Poldrack

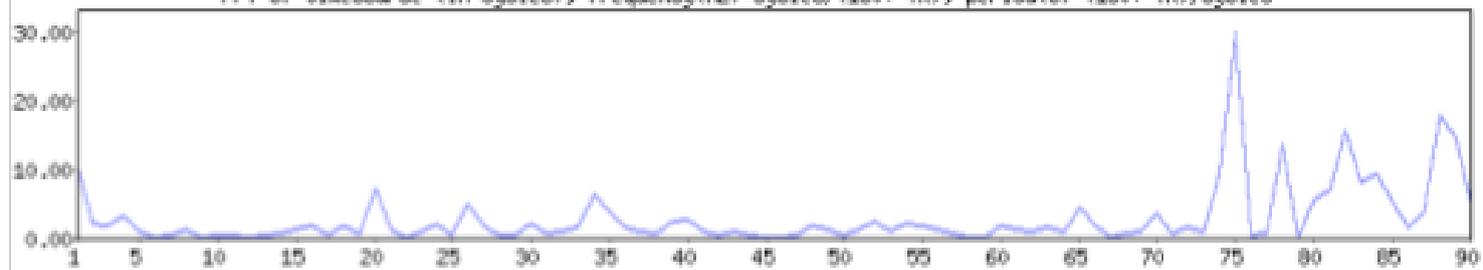
High frequency noise



Timecourse (in TRs)

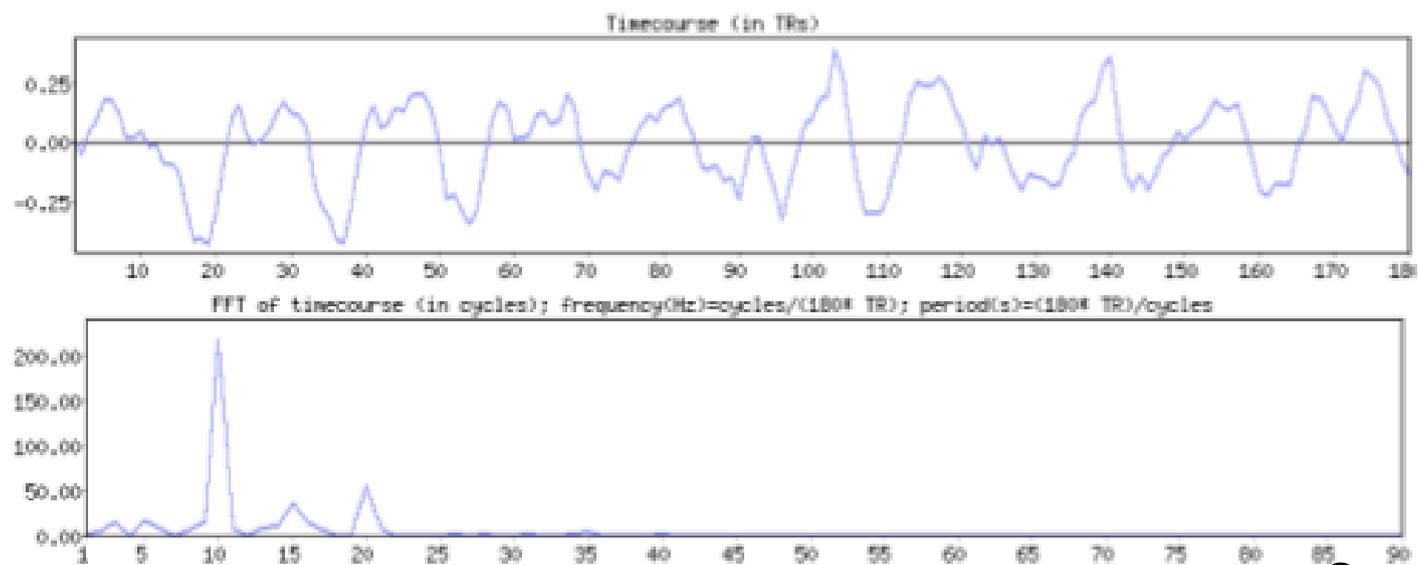
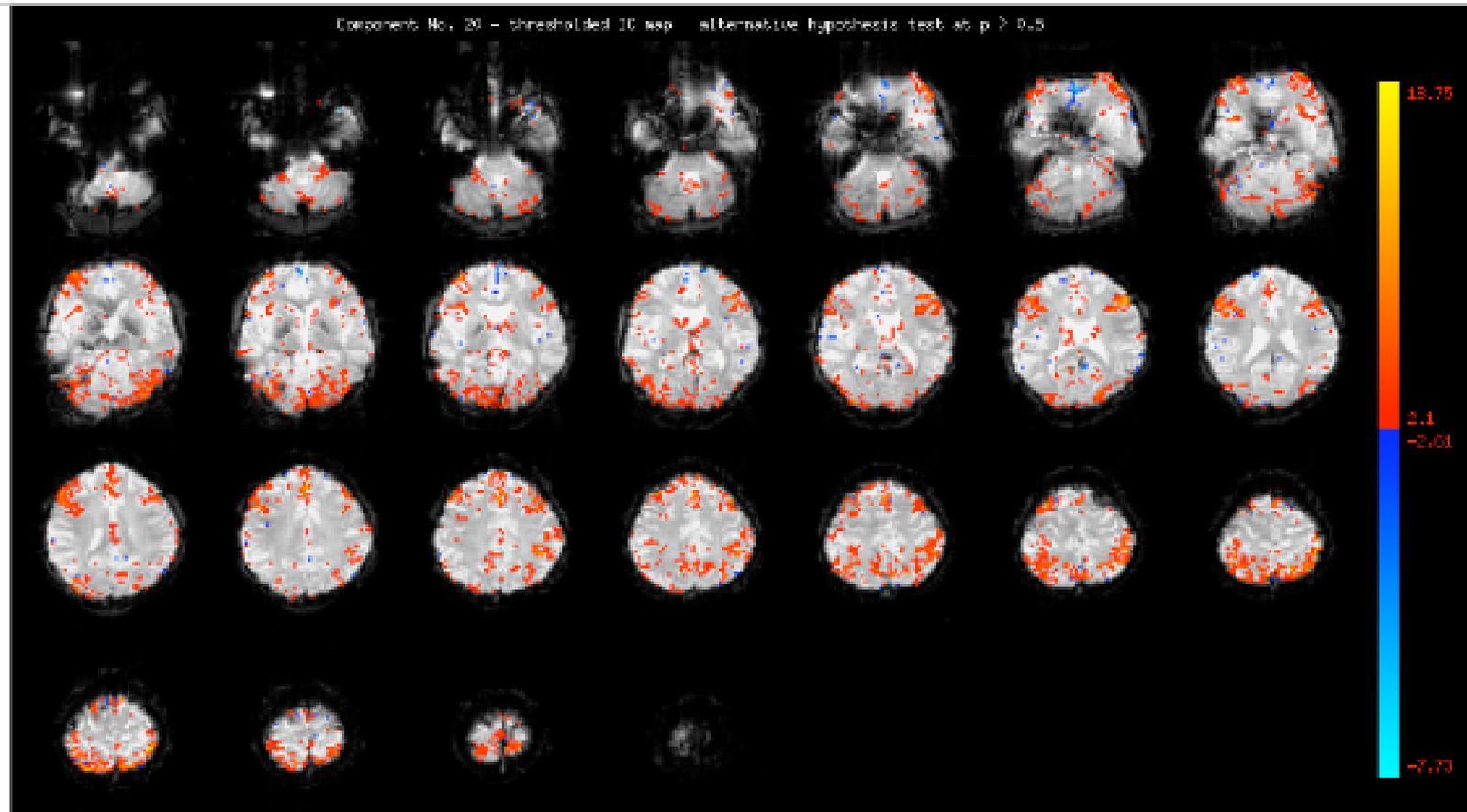


FFT of timecourse (in cycles): frequency(Hz)=cycles/(180* TR); period(s)=(180* TR)/cycles



Source: R. Poldrack

Task



Registration

Functional

Anatomical

Template

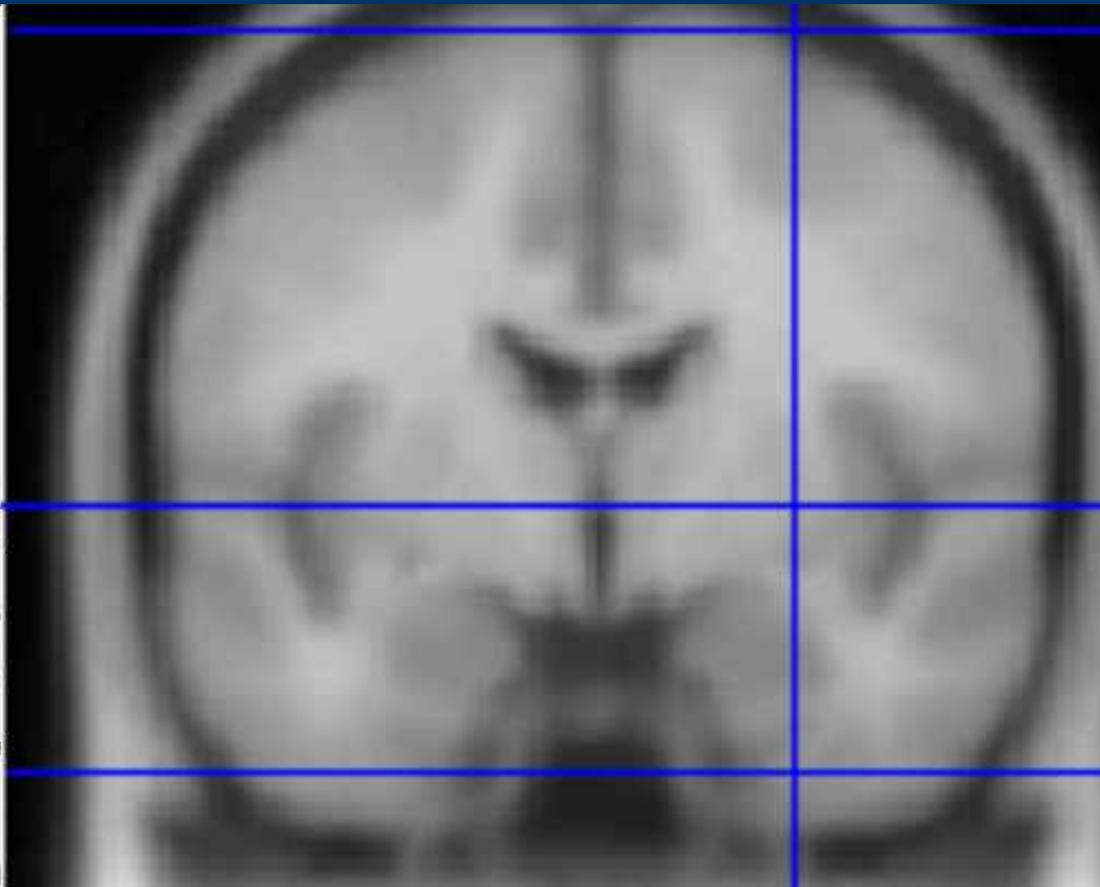
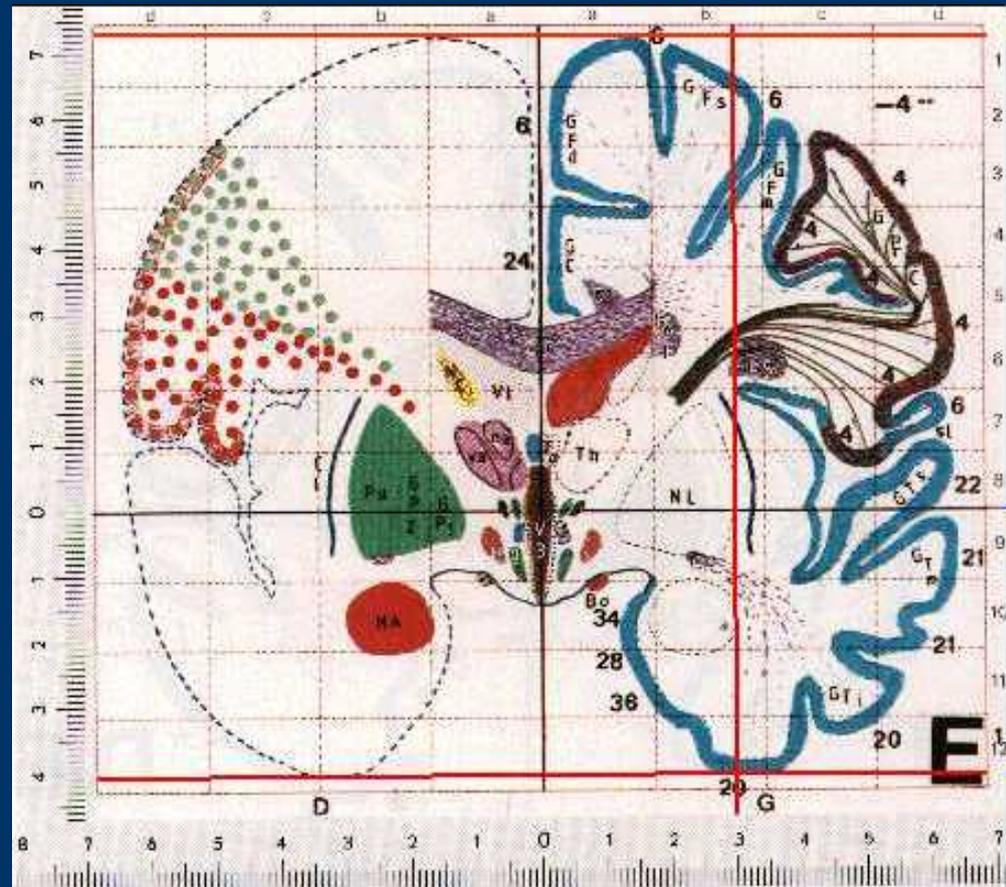


Standard Space

- Common reference frame
- Talairach & Tournoux 1988, based on post mortem dissection of 1 brain
- MNI (152) non linear average of multiple individuals



Standard Space



Registration

- 1. Transformation:** How to manipulate an image to fit it from its native space into a different space?
- 2. Cost function:** How to assess the quality of the manipulation?
- 3. Interpolation:** How create the intensity values to be assigned to the new “grid”?

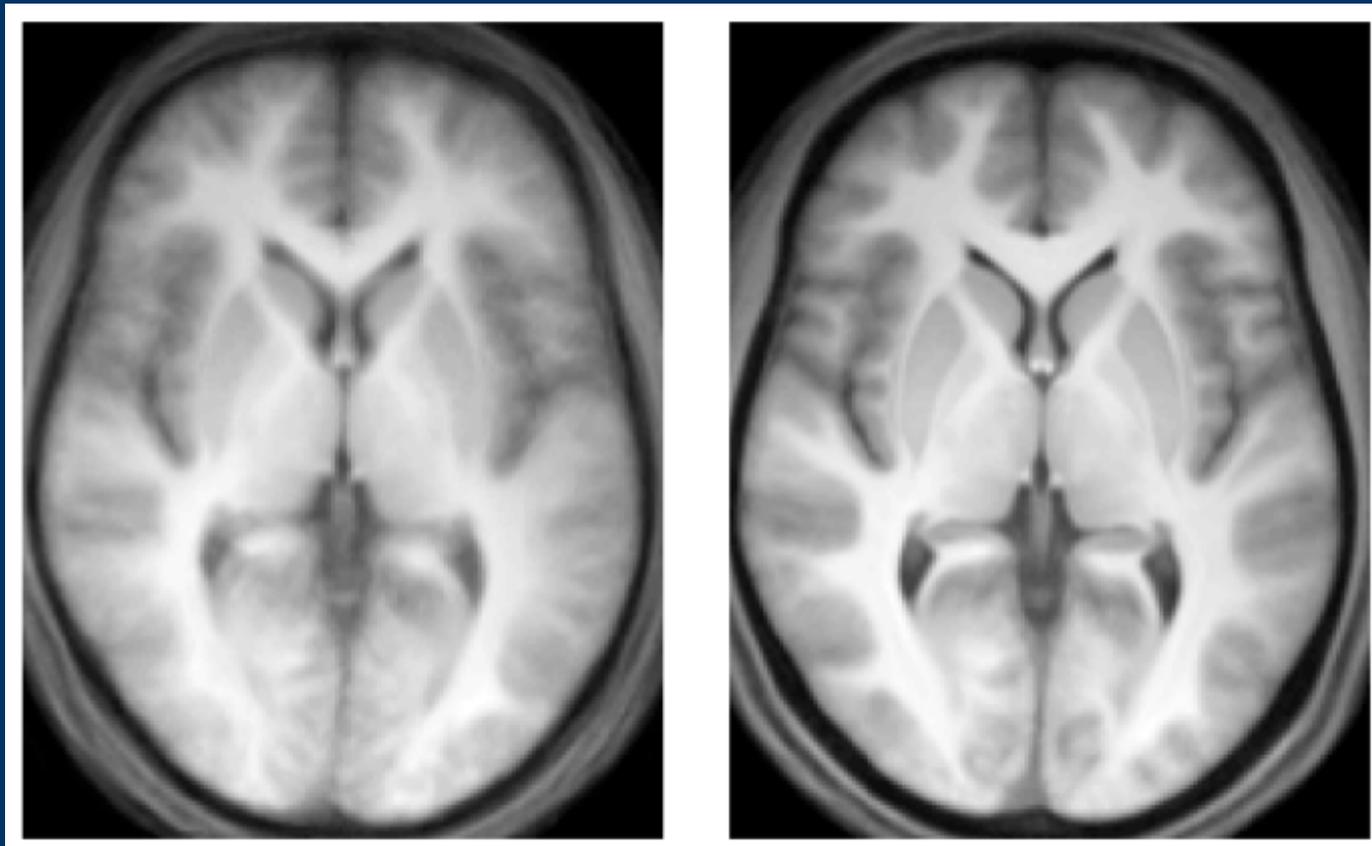


Transformations

- Rigid body (6dof):
 - 3 rotations, 3 translations
 - Typically used for intra-subject registration.
 - Rigid body + global scaling (7dof)
 - 3 rotations, 3 translations, global scaling
 - Typically used for within subject/between modalities (i.e., functional to structural)
 - Affine (12dof)
 - 3 rotations, 3 translations
 - 3 scalings, 3 shears/skews
 - Typically used for registering a subject to the template
-
-

Transformations

- Non-linear (> 12 dof)
 - Can be local
 - Can be constrained (e.g., regularization, topology preservation)



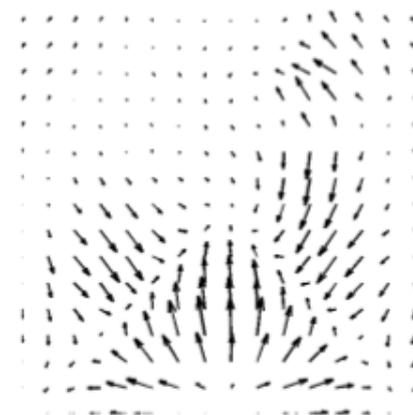
Transformations

- Non-linear (> 12dof)
 - Can be local
 - Can be constrained (e.g., regularization, topology preservation)

$$A = \begin{pmatrix} a_{11} & a_{12} & a_{13} & a_{14} \\ a_{21} & a_{22} & a_{23} & a_{24} \\ a_{31} & a_{32} & a_{33} & a_{34} \\ 0 & 0 & 0 & 1 \end{pmatrix}$$

An affine transformation is represented by these 12 numbers.

This matrix multiplies coordinate vectors to define the transformed coordinates.



A non-linear transformation is represented by a **deformation field**.

Cost functions



FLIRT: Cost Functions

Important: Allowable image modalities

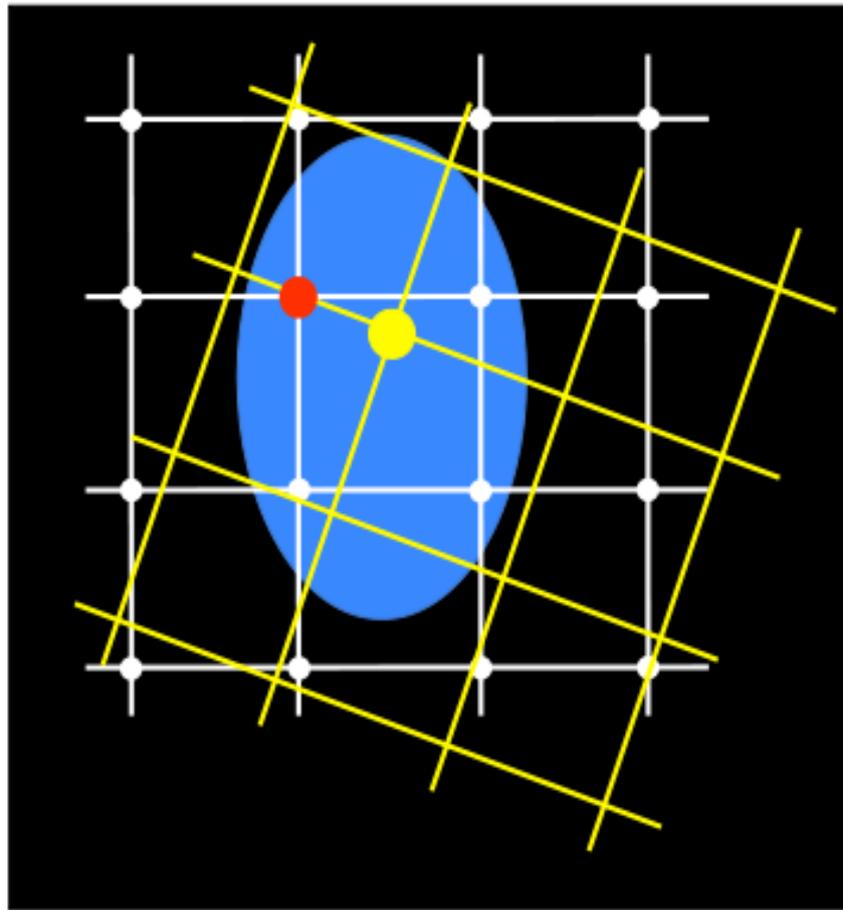
Less important: Details

Least Squares	Same modality (exact sequence parameters)
Normalised Correlation	Same modality (can change brightness & contrast)
Correlation Ratio	Any MR modalities
Mutual Information	Any modalities (including CT, PET, etc.)
Normalised Mutual Info.	Any modalities (including CT, PET, etc.)



Interpolation

Finds intensity values between grid points



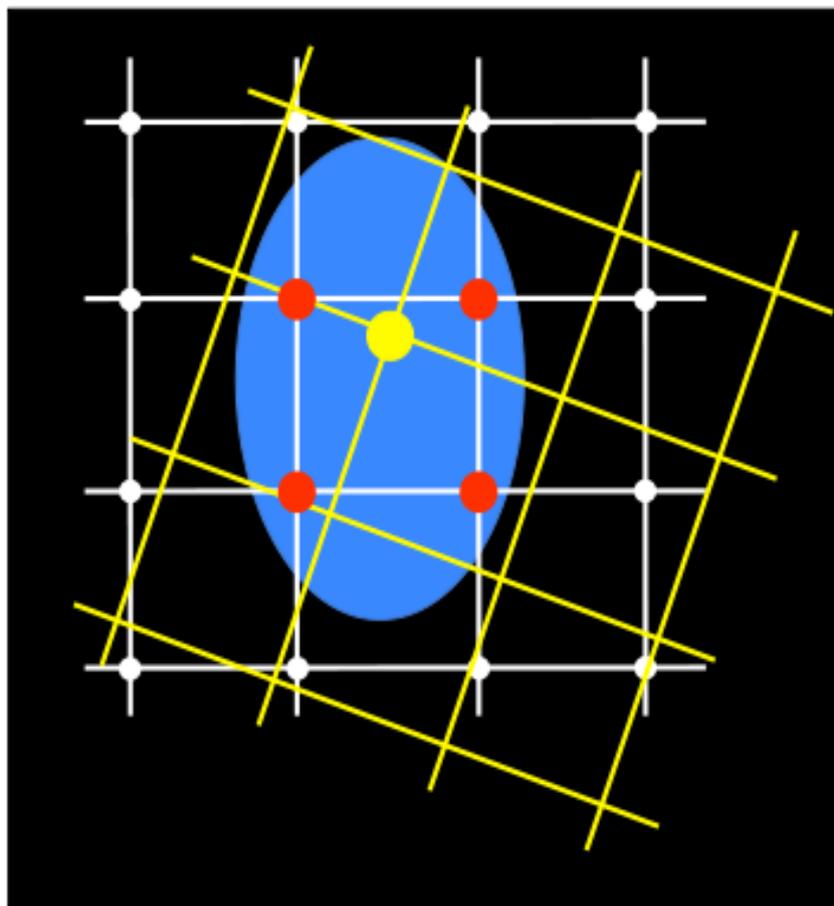
Various types include

- Nearest Neighbour
- Trilinear
- Sinc
- Spline
- k-Space methods



Interpolation

Finds intensity values between grid points



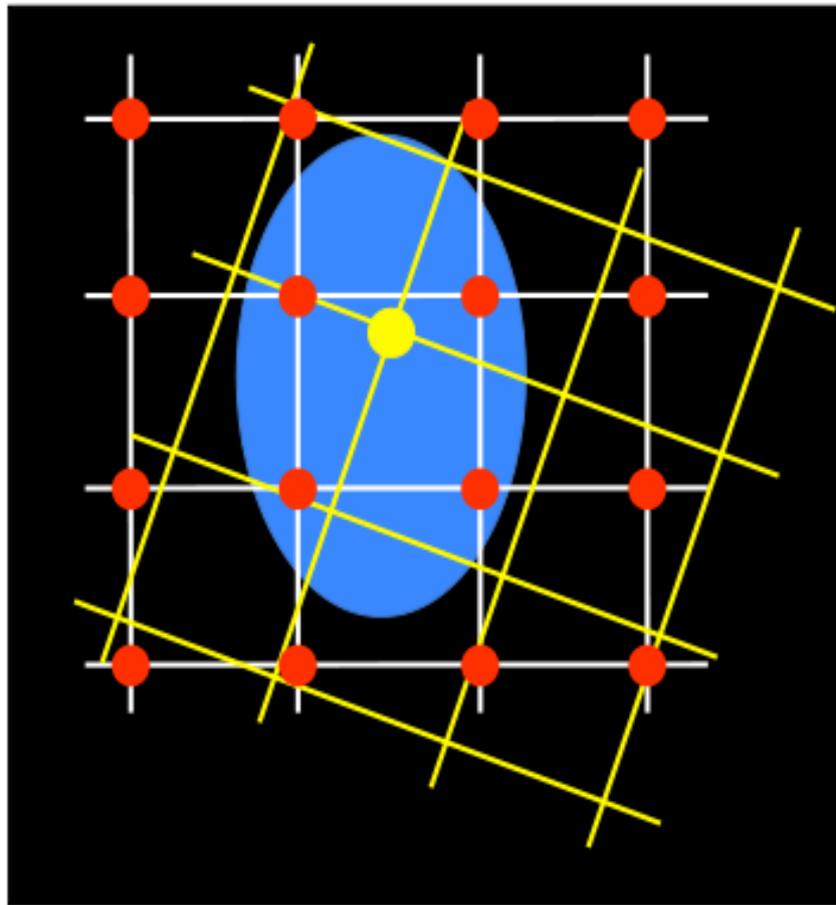
Various types include

- Nearest Neighbour
- **Trilinear**
- Sinc
- Spline
- k-Space methods



Interpolation

Finds intensity values between grid points



Various types include

- Nearest Neighbour
- Trilinear
- Sinc
- Spline
- k-Space methods

Considerations: speed, accuracy, stability

