

The logo features a large, stylized 'NP' in a golden-yellow color with a blue glow effect. To the right of the 'NP', the words 'NEUROIMAGING', 'TRAINING', and 'PROGRAM' are stacked vertically in a bold, black, sans-serif font.

# **NP NEUROIMAGING TRAINING PROGRAM**

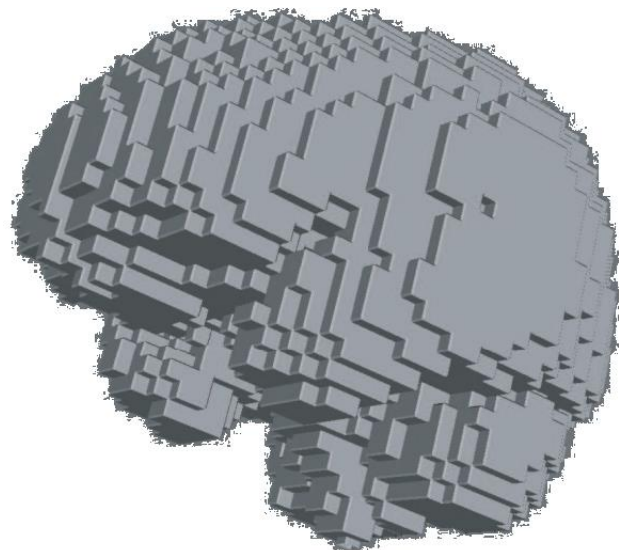
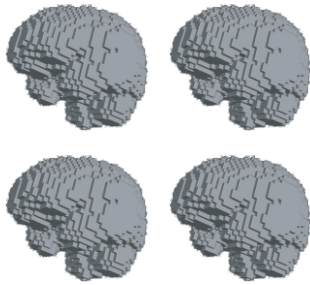


# PREPROCESSING

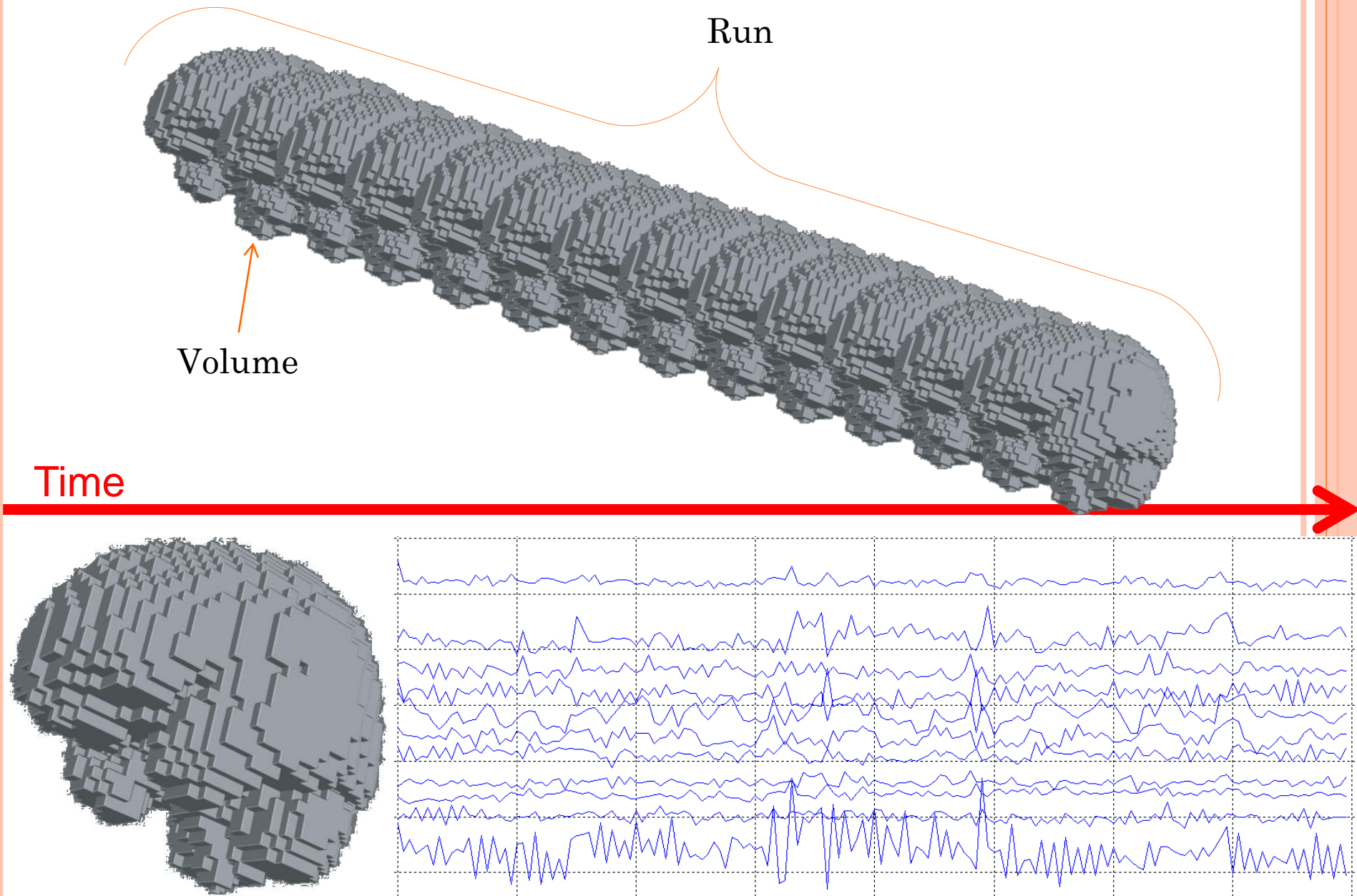
**Martin M. Monti**  
**UCLA Psychology**

**NITP 2015**

# TYPICAL DATASET



# TYPICAL DATASET



# TYPICAL FMRI ANALYSIS SEQUENCE

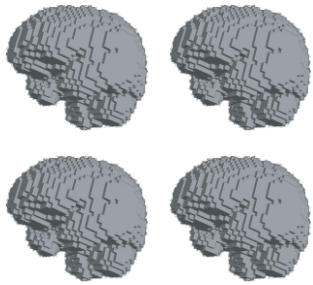
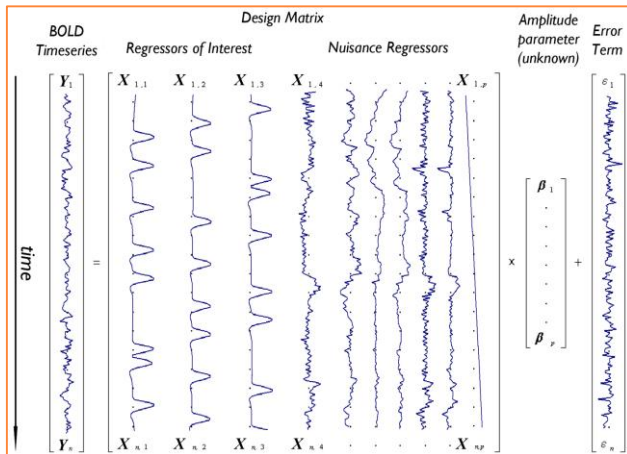
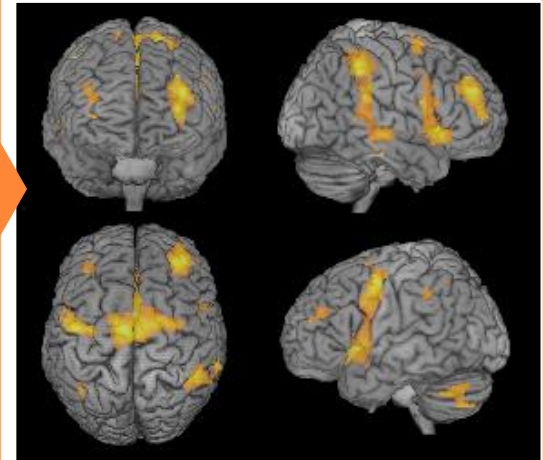


Image Pre-processing

## Single Subject Analysis



## Group Analysis



# PREPROCESSING: WHAT/WHY?

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



# SOURCES OF NOISE IN FMRI

## 1. Hardware & acquisition related:

- Thermal noise (intrinsic noise)

- System noise

- Field inhomogeneities

- Slice acquisition timing

## 2. Subject related

- Oscillatory physiological noise (heartbeat, respiration)

- Field inhomogeneities

- Head motion

- Psychological (alertness, learning)

## 3. White noise



# CORRECTING FOR NOISE IN FMRI

1. Before scanning (maximize SNR):
  - i. Choose good technology (field strength, coils, ...)
  - ii. Choose good sequence (TE, voxel size, ...)
  - iii. Be informed about the healthy of your scanner (QA)
2. After scanning (detect & correct):
  - i. Look at your data (i.e., data quality check)
  - ii. Look at your data (again and again)
  - iii. Pre-processing (“standard”, ICA)
  - iv. Re-look at your data

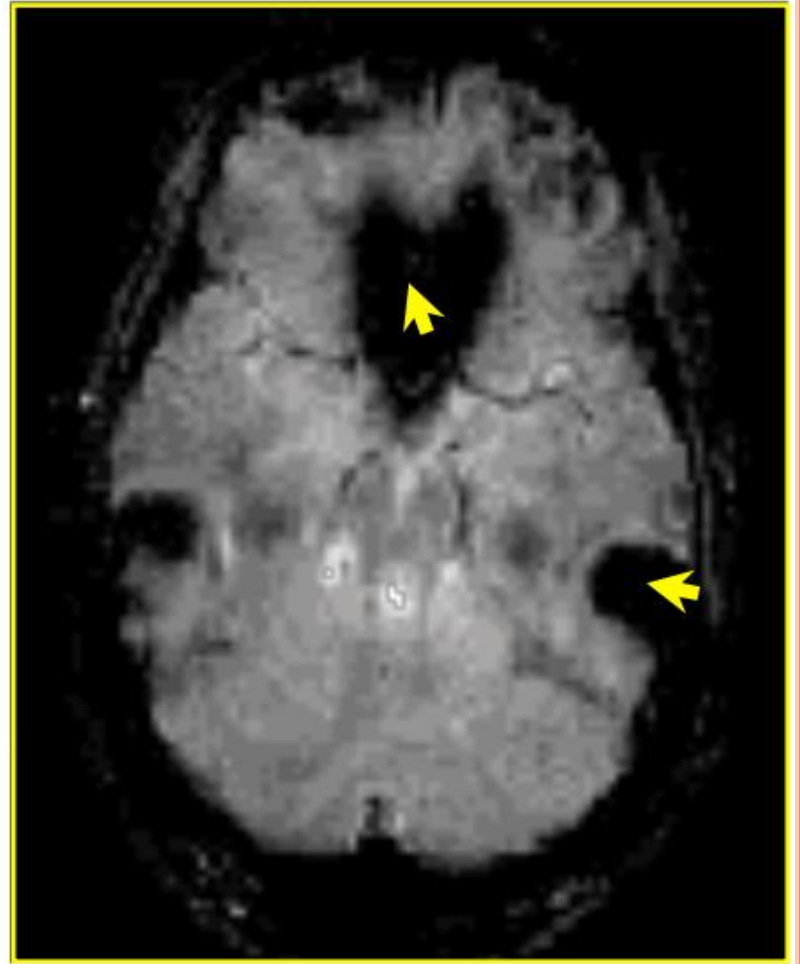




# SEQUENCE PARAMETERS: TE



Short TE



Long TE

S Clare

# CORRECTING FOR NOISE IN FMRI

1. Before scanning (maximize SNR):
  - i. Choose good technology (field strength, coils, ...)
  - ii. Choose good sequence (TE, voxel size, ...)
  - iii. Be informed about the healthy of your scanner (QA)
2. After scanning (detect & correct):
  - i. Look at your data (i.e., data quality check)
  - ii. Look at your data (again and again)
  - iii. Pre-processing (“standard”, ICA)
  - iv. Re-look at your data



# PREPROCESSING: WHAT/WHY?

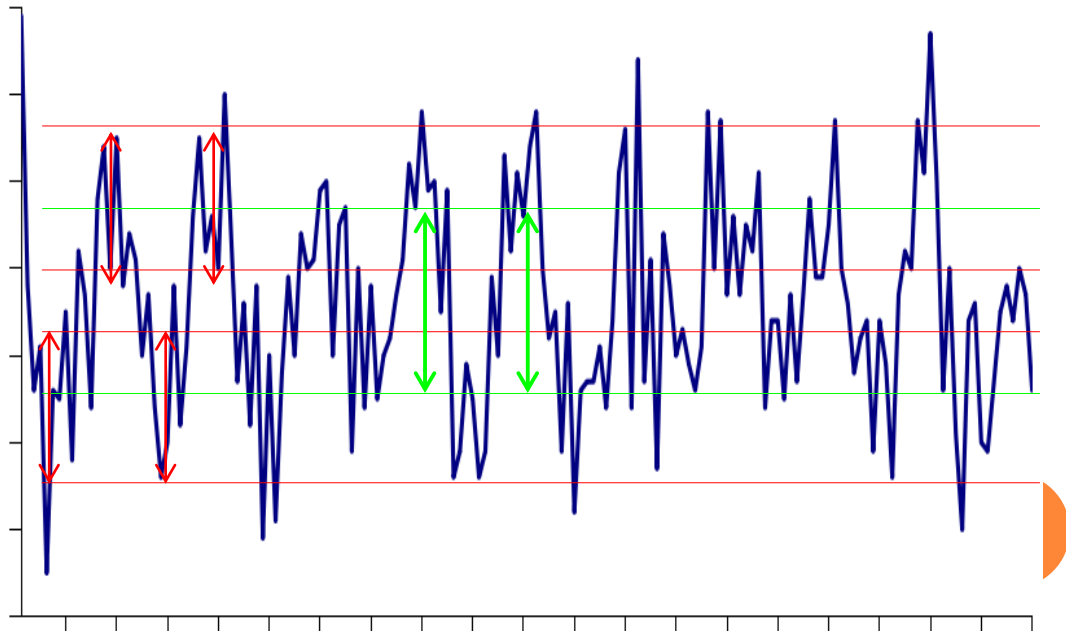
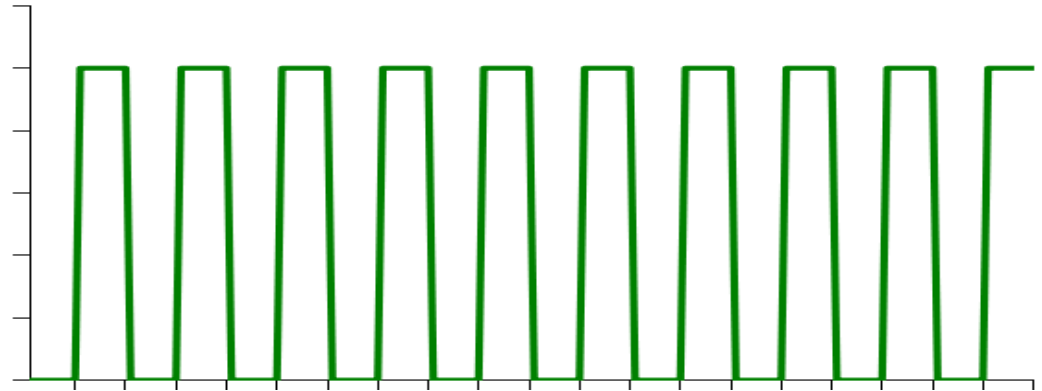
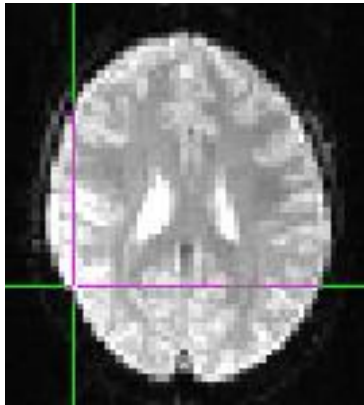
Preprocessing is a series of data transformations (“data conditioning”) aimed at reducing sources of noise in order to:

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



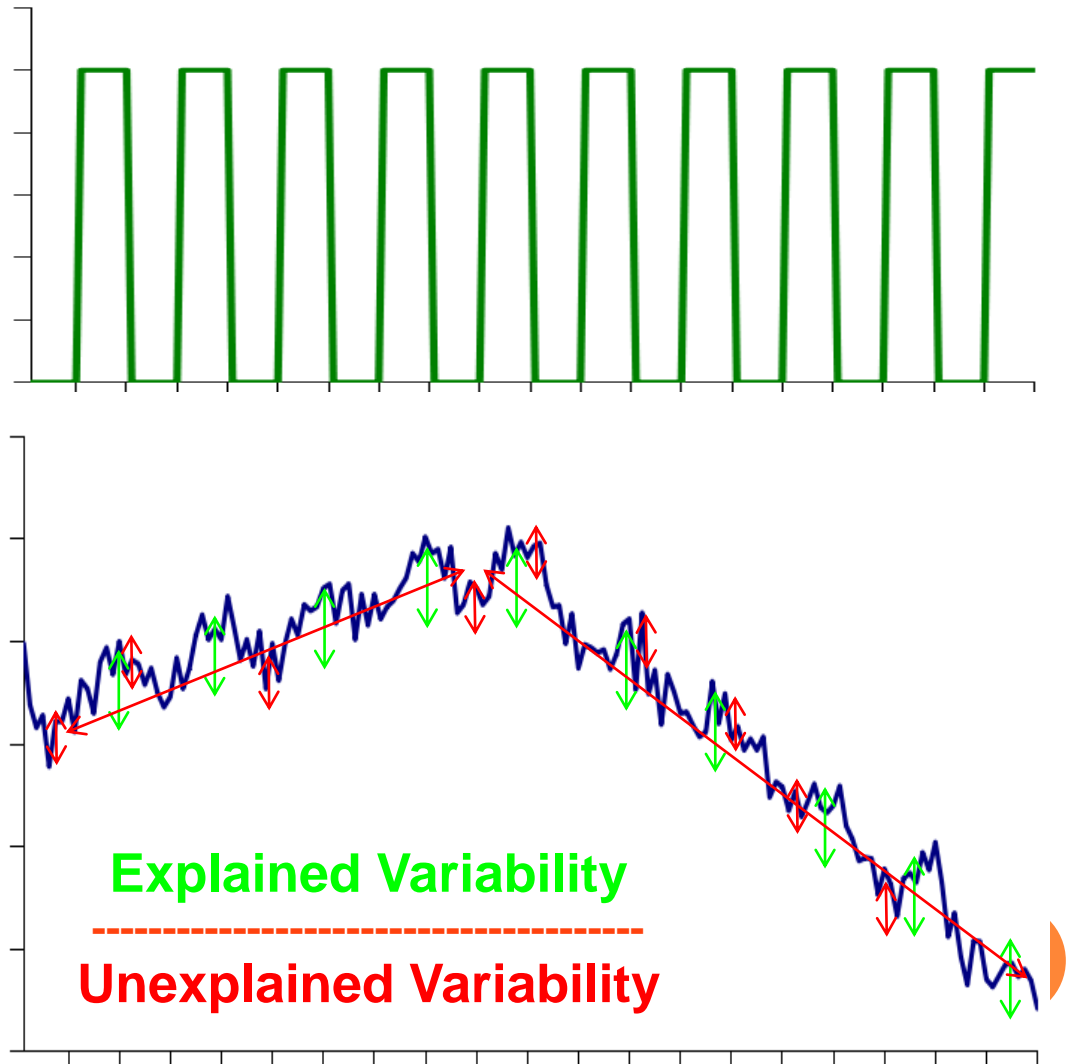
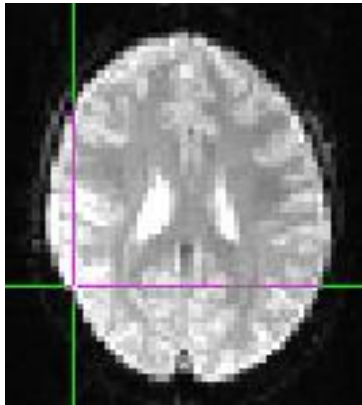
# SAMPLE EXPERIMENT: SNR

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



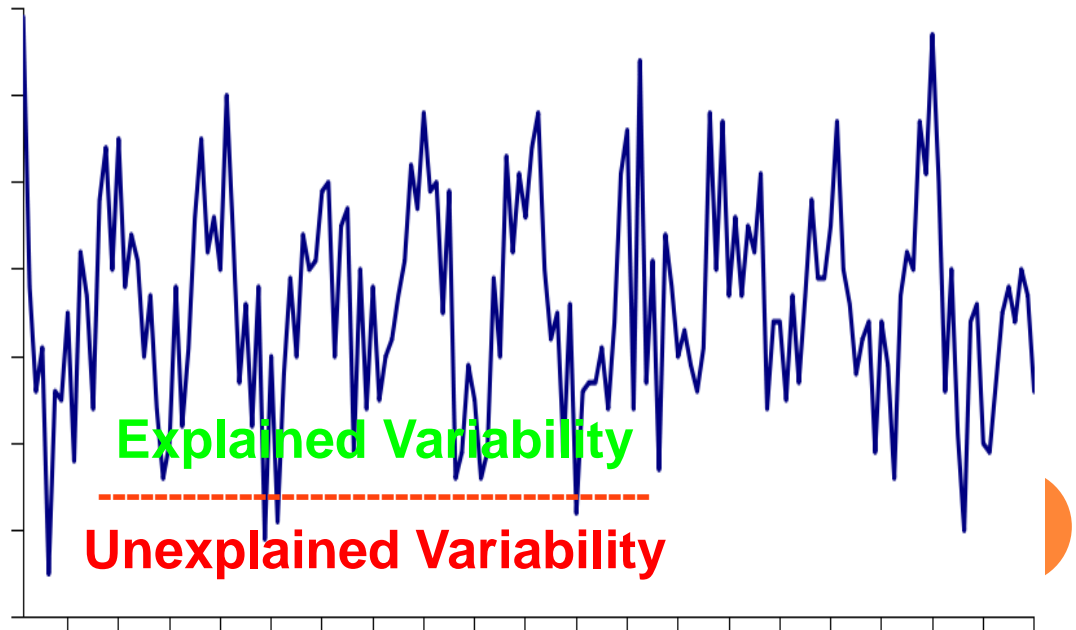
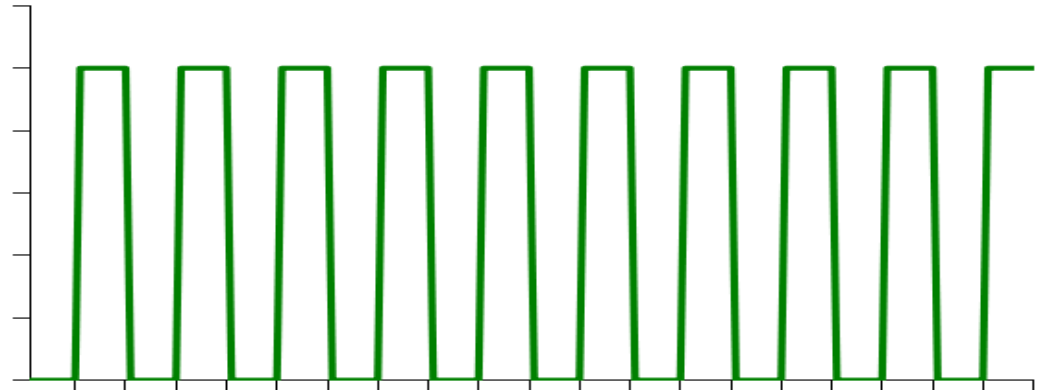
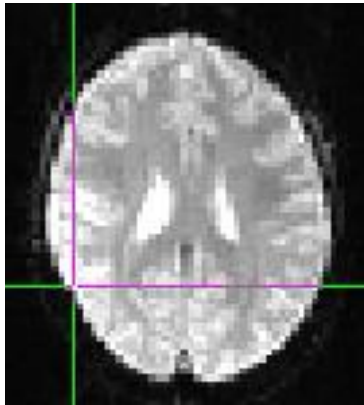
# SAMPLE EXPERIMENT: SNR

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



# SAMPLE EXPERIMENT: SNR

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



# THE GENERAL LINEAR MODEL (GLM)

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

fMRI Signal	Design Matrix	Parameter	Residuals
<i>“our data”</i>	<i>“what we CAN explain”</i>	<i>“how much of it we CAN explain”</i>	<i>“what we CANNOT explain”</i>
=	x	+	



# PREPROCESSING

- i. Motion correction
- ii. Slice timing correction
- iii. Spatial filtering
- iv. Temporal filtering
- v. Intensity normalization



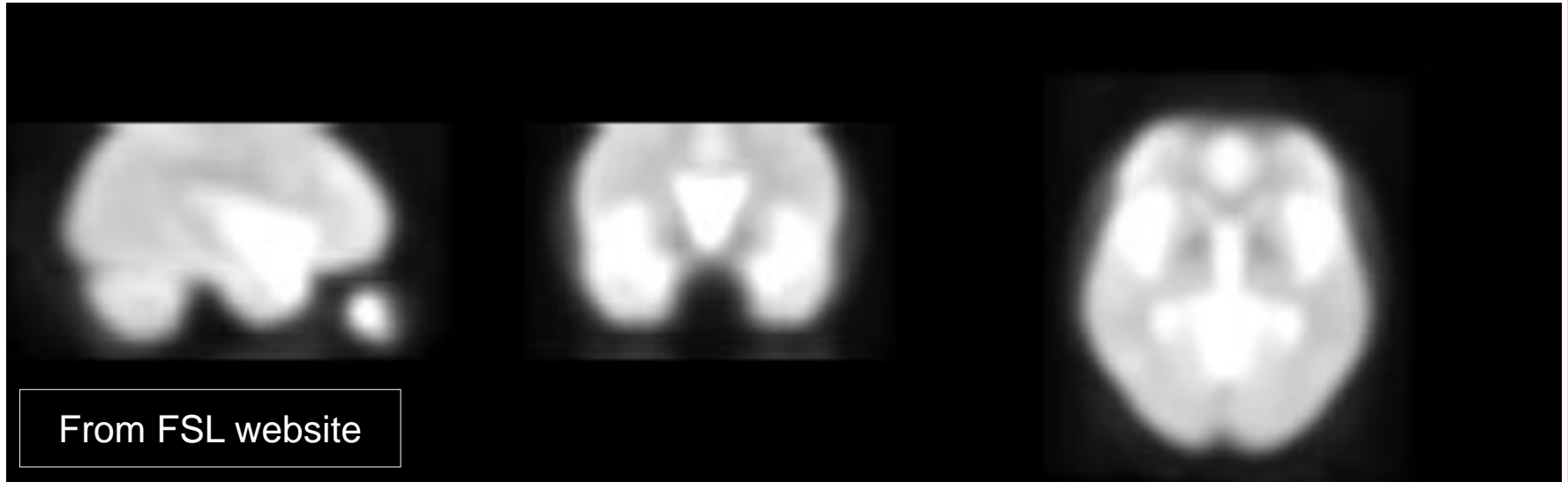


# PREPROCESSING

- i. Motion correction
- ii. Slice timing correction
- iii. Spatial filtering
- iv. Temporal filtering
- v. Intensity normalization



# SUBJECT MOTION



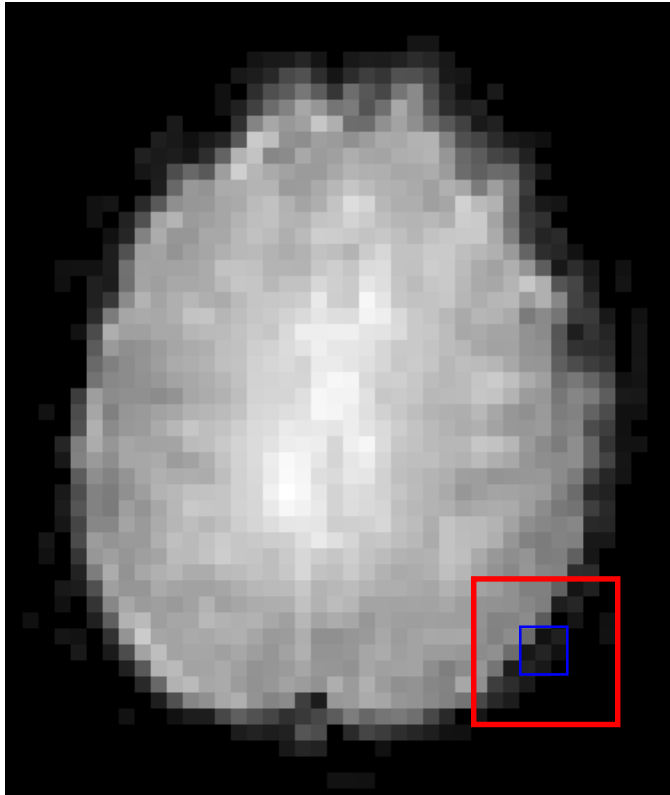
Motion within a time-series can have several unwanted consequences:

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

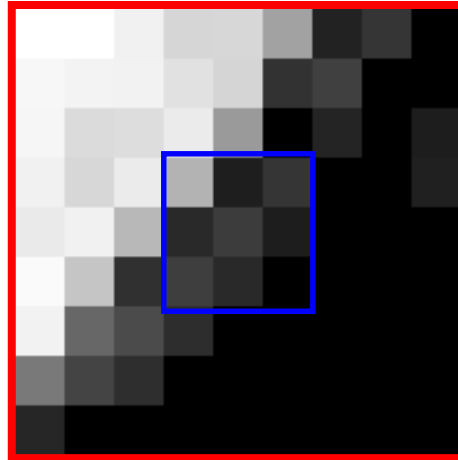


# SUBJECT MOTION

A

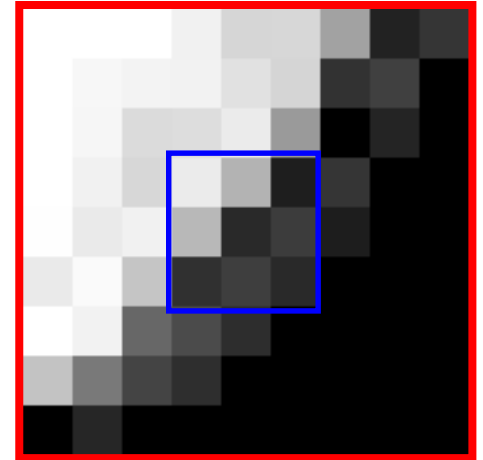


B



507	89	154
119	171	83
179	117	53

C

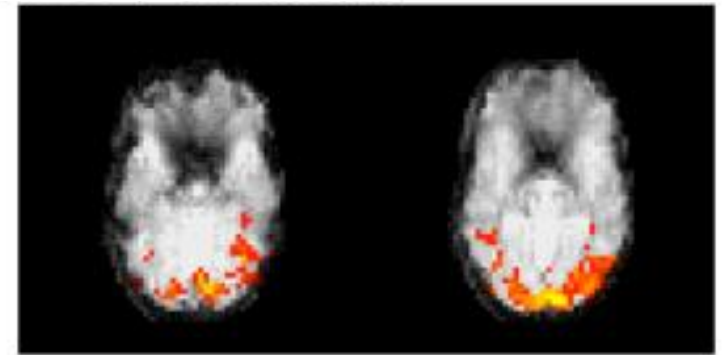
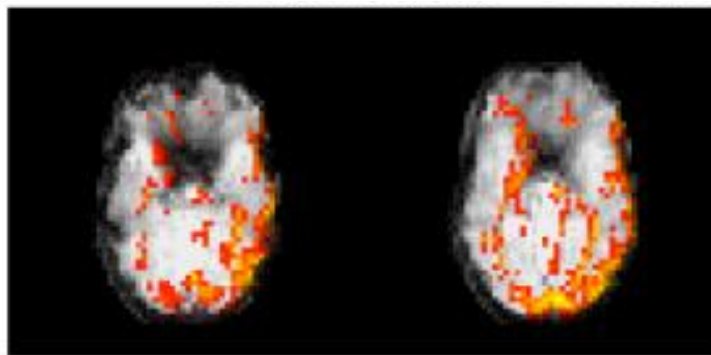
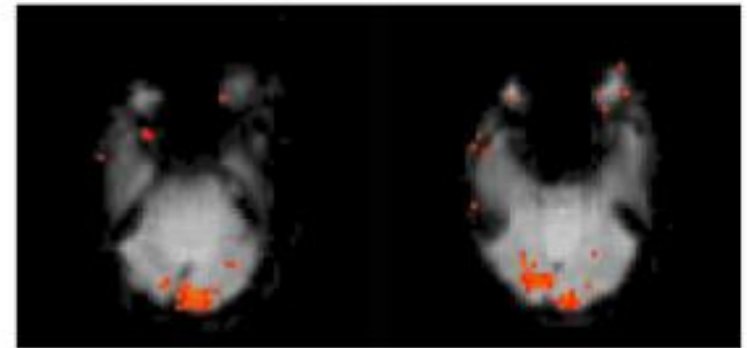
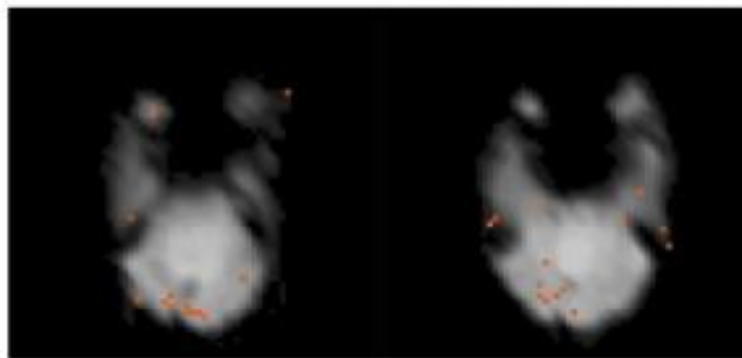


663	507	89
520	119	171
137	179	117

Huettel et al. Functional Magnetic Resonance Imaging



# Effect of Motion Correction



Without MC

With MC

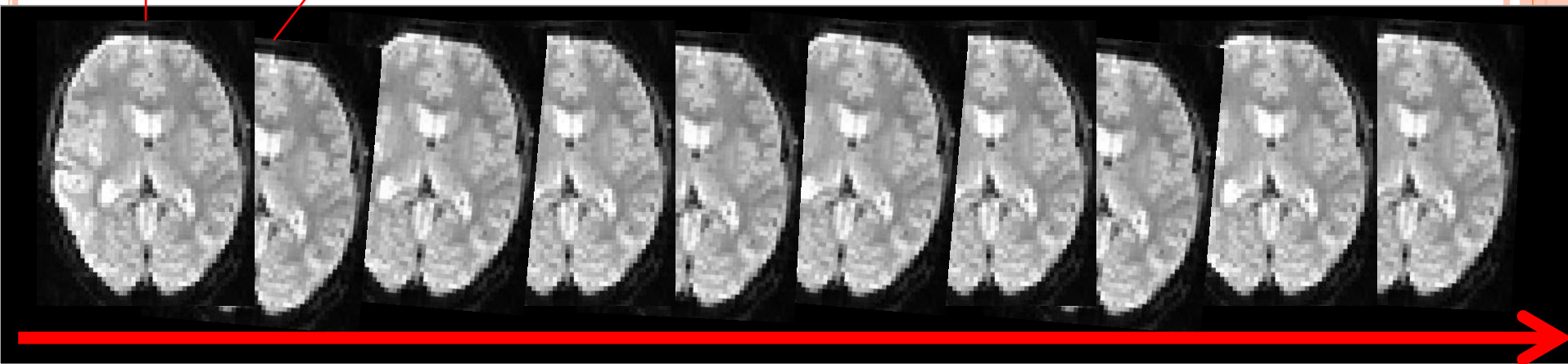
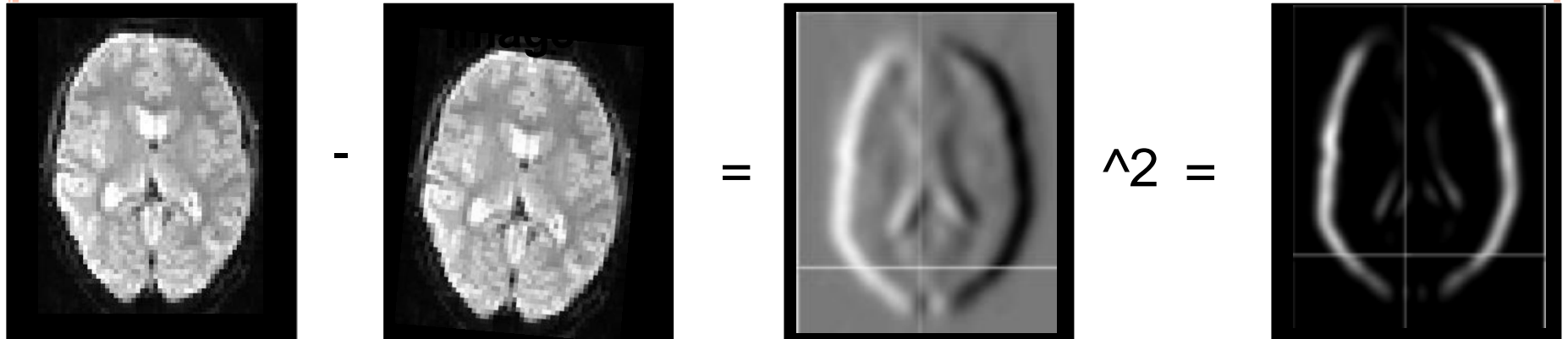
# MOTION CORRECTION

**Reference**

***lth***

**Difference**

**Variance**



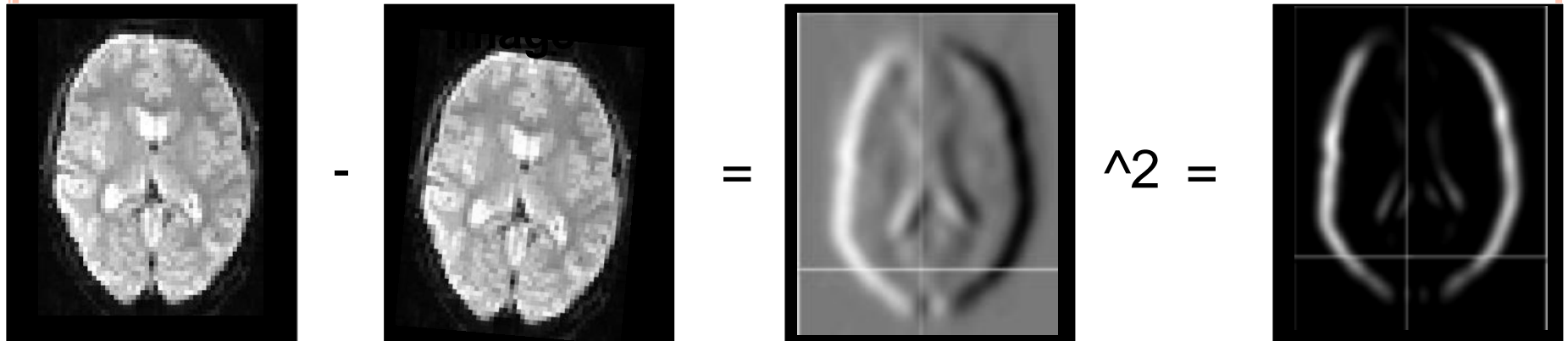
# MOTION CORRECTION

**Reference**

***l*th**

**Difference**

**Variance**



Rigid body (6dof)

Rigid body transformations parameterised by:

**Translations**

**Pitch**

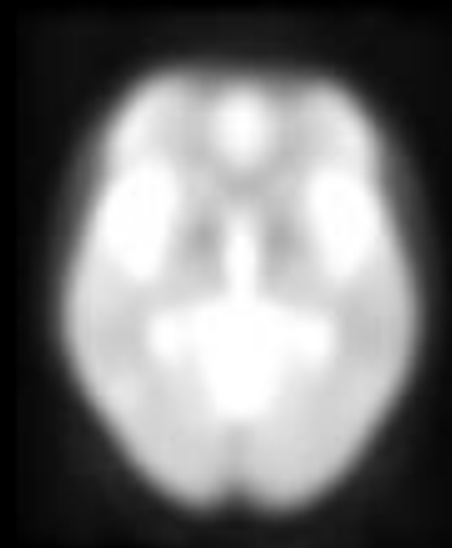
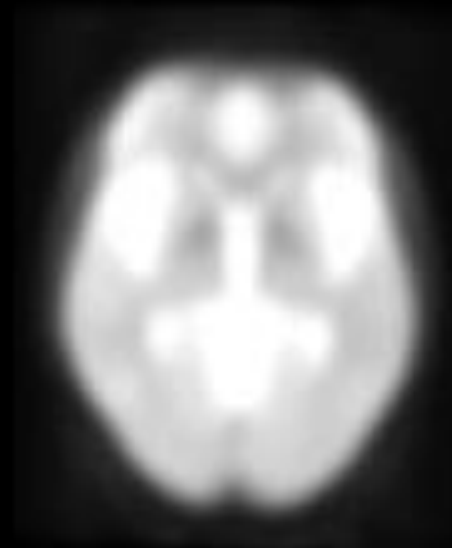
**Roll**

**Yaw**

$$\begin{pmatrix} 1 & 0 & 0 & X_{trans} \\ 0 & 1 & 0 & Y_{trans} \\ 0 & 0 & 1 & Z_{trans} \\ 0 & 0 & 0 & 1 \end{pmatrix} \times \begin{pmatrix} 1 & 0 & 0 & 0 \\ 0 & \cos(\Phi) & \sin(\Phi) & 0 \\ 0 & -\sin(\Phi) & \cos(\Phi) & 0 \\ 0 & 0 & 0 & 1 \end{pmatrix} \times \begin{pmatrix} \cos(\Theta) & 0 & \sin(\Theta) & 0 \\ 0 & 1 & 0 & 0 \\ -\sin(\Theta) & 0 & \cos(\Theta) & 0 \\ 0 & 0 & 0 & 1 \end{pmatrix} \times \begin{pmatrix} \cos(\Omega) & \sin(\Omega) & 0 & 0 \\ -\sin(\Omega) & \cos(\Omega) & 0 & 0 \\ 0 & 0 & 1 & 0 \\ 0 & 0 & 0 & 1 \end{pmatrix}$$

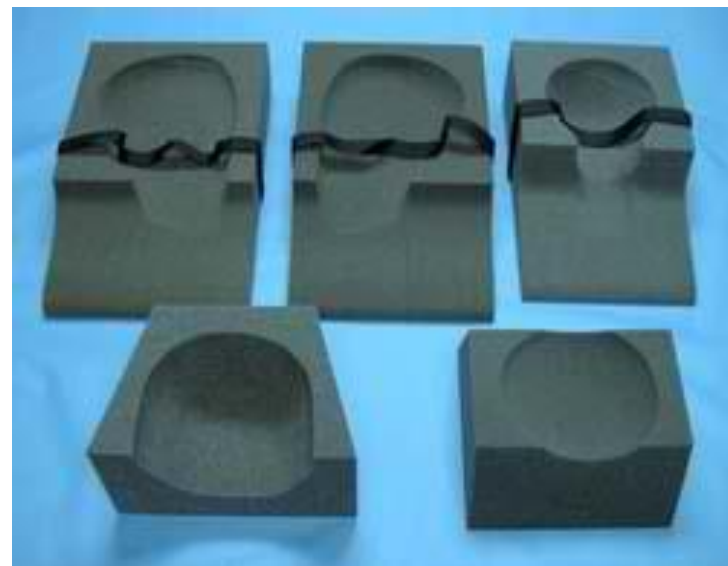
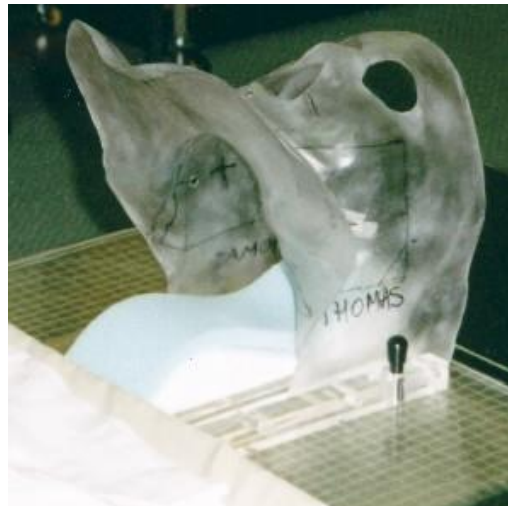
# VIEWING MOTION CORRECTION



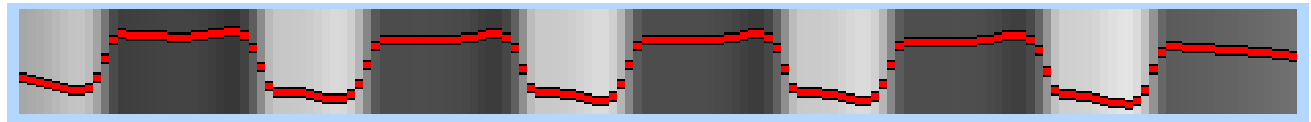
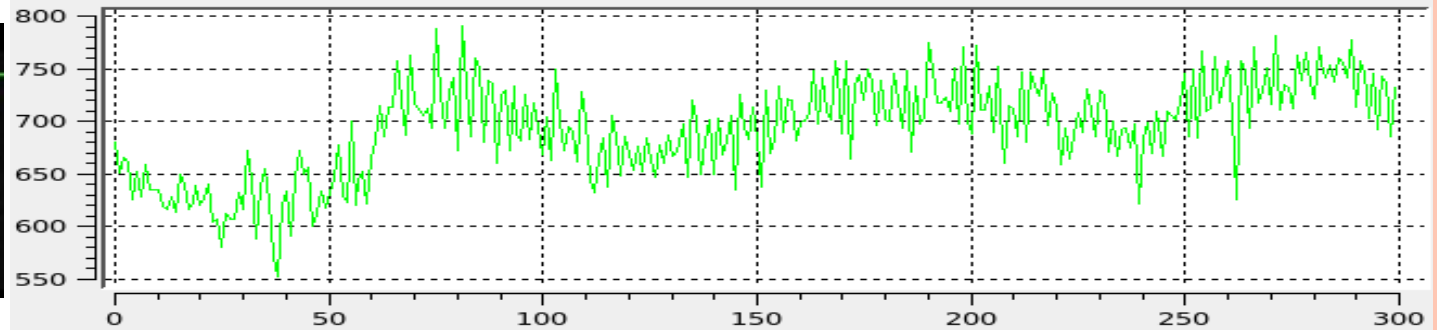
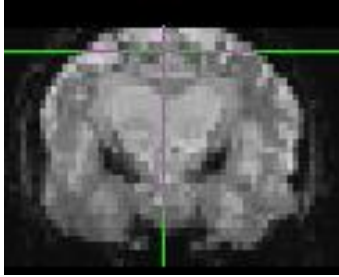




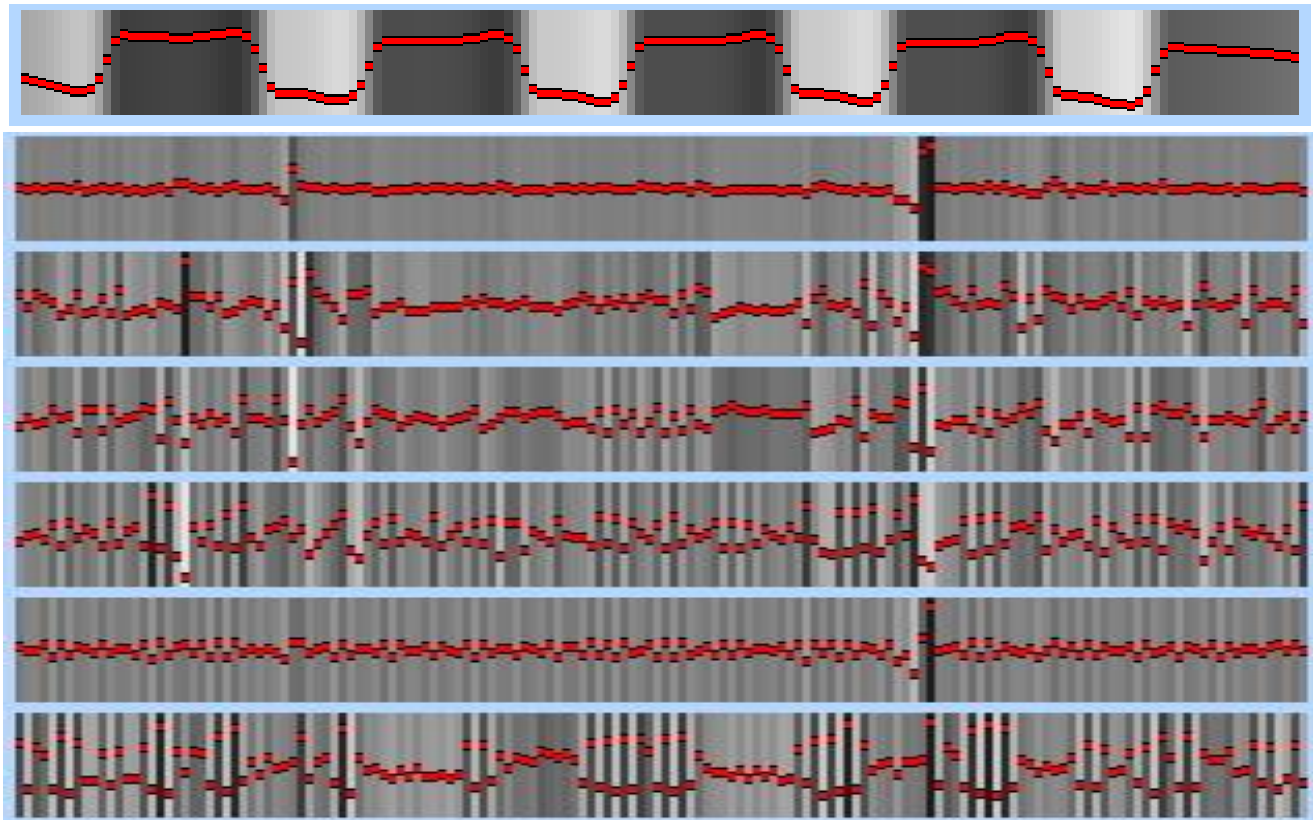
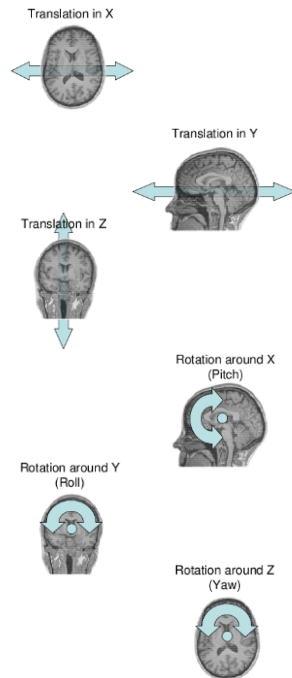
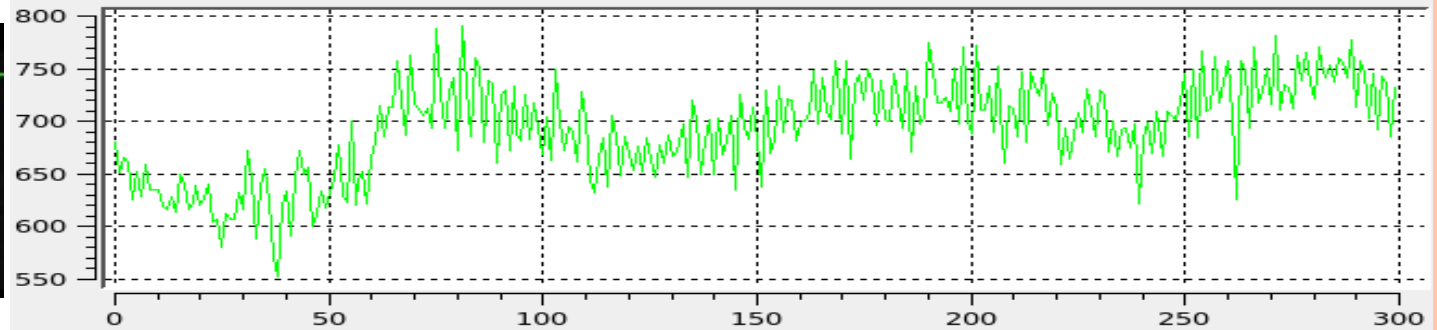
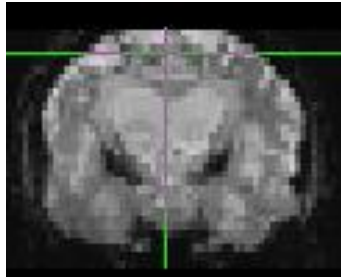
# COPING WITH MOTION I: PREVENT IT



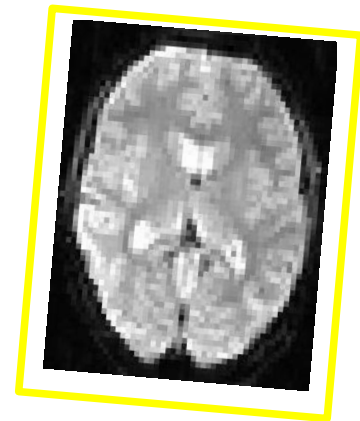
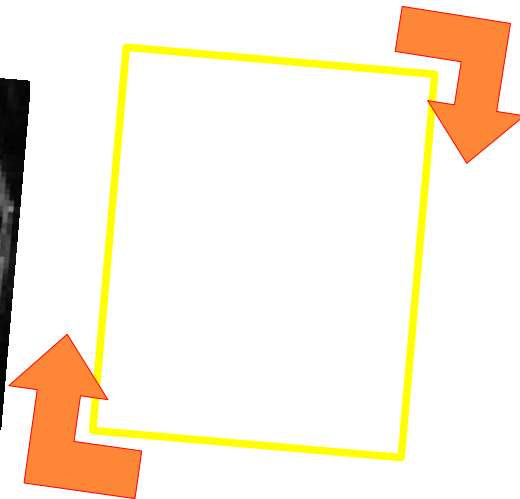
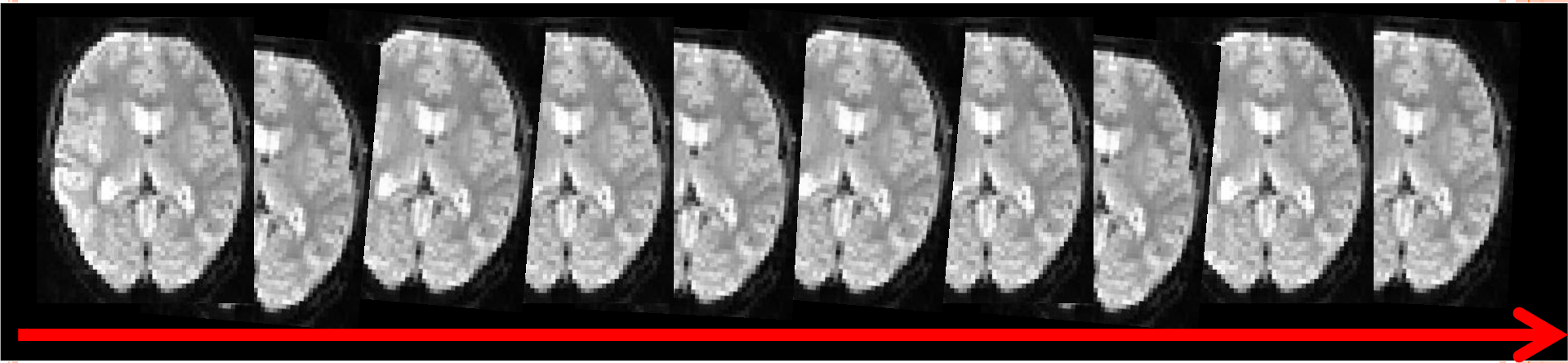
# COPING WITH MOTION II(A): MODEL IT



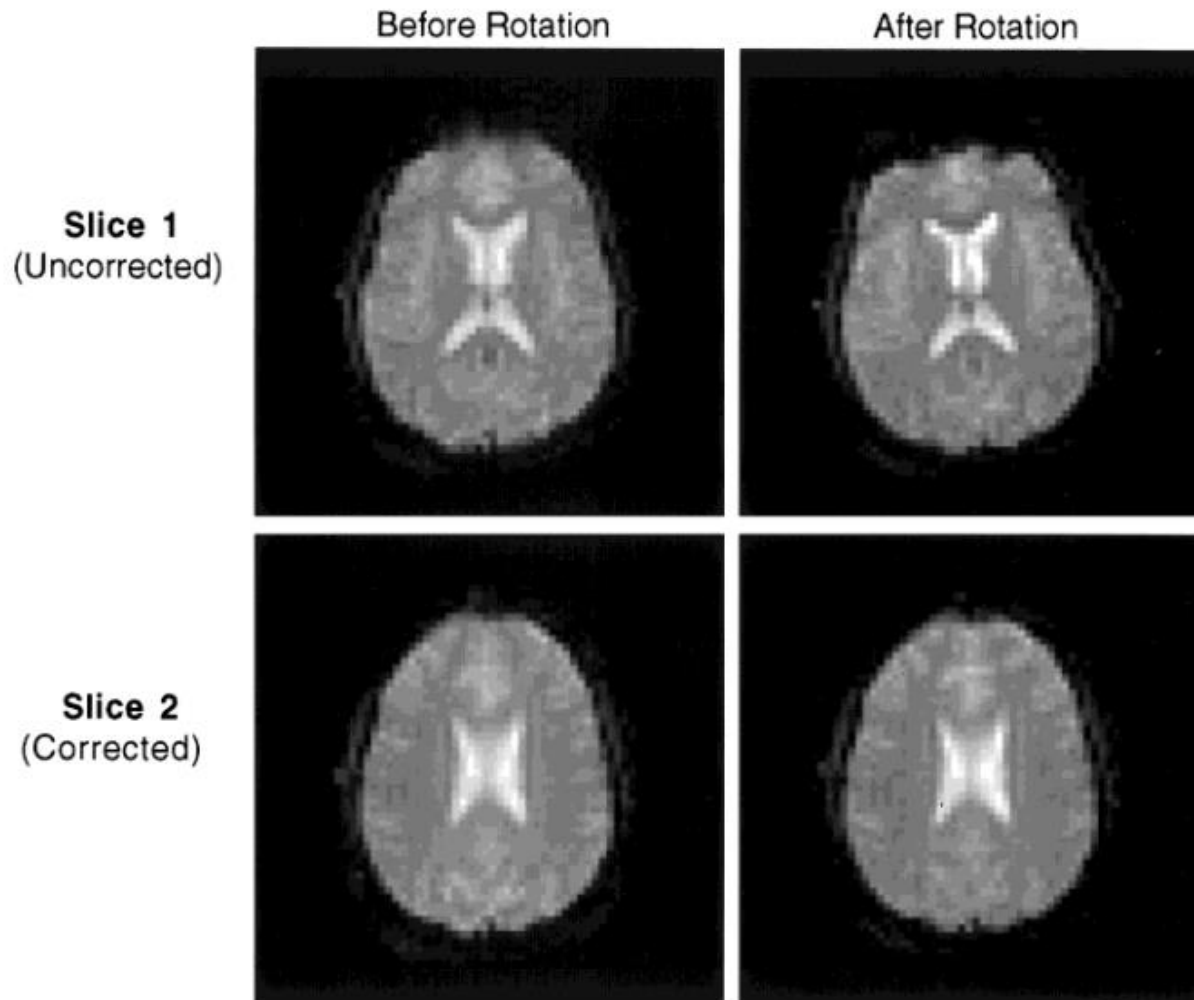
# COPING WITH MOTION II(B): MODEL IT



# COPING WITH MOTION III: PROSPECTIVE MC



# PROSPECTIVE MOTION CORRECTION



# 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\*
  - 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

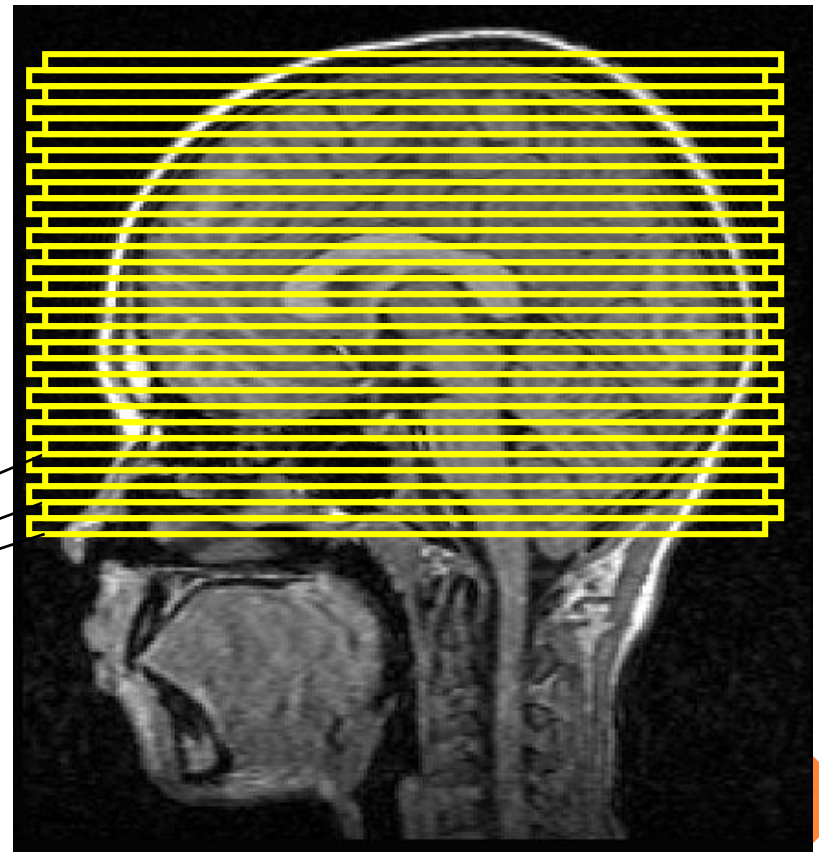
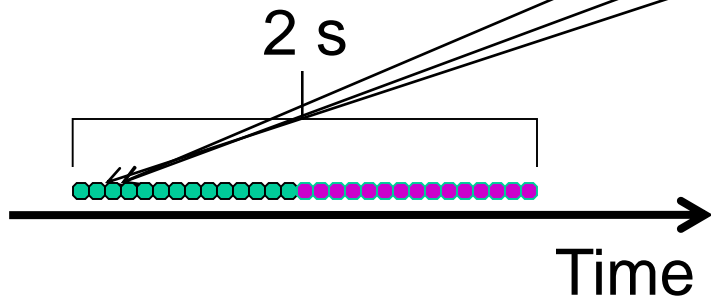
- i. Motion correction
- ii. Slice timing correction**
- iii. Spatial filtering
- iv. Temporal filtering
- v. Intensity normalization





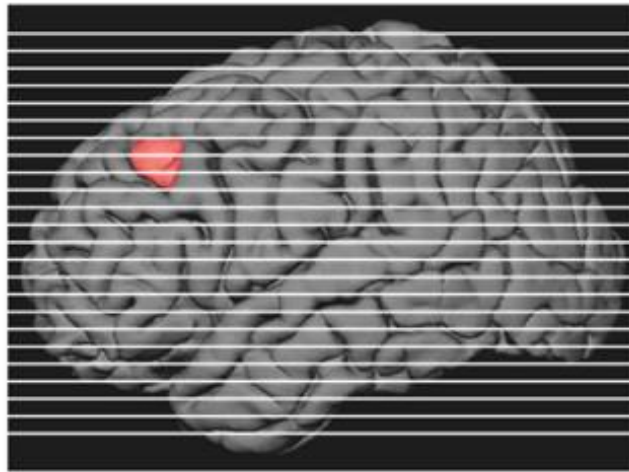
# SLICE TIMING CORRECTION

- 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).

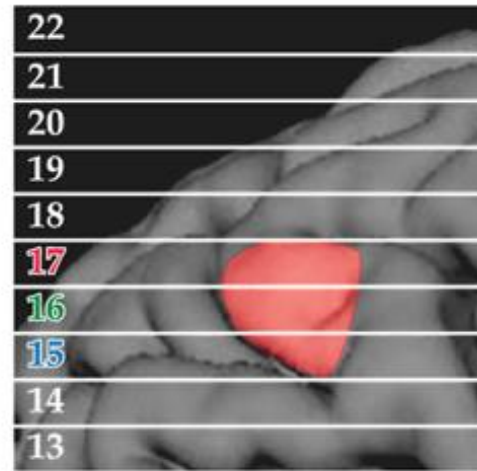


# SLICE TIMING CORRECTION

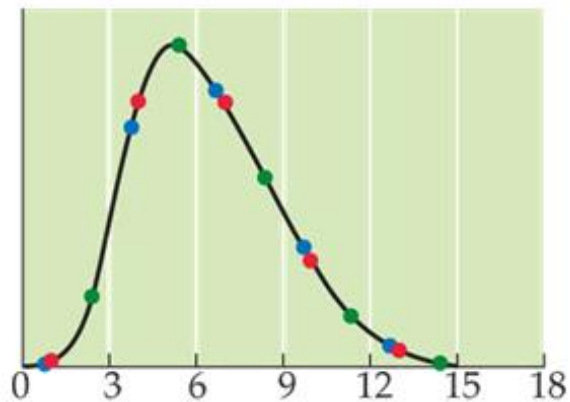
(A)



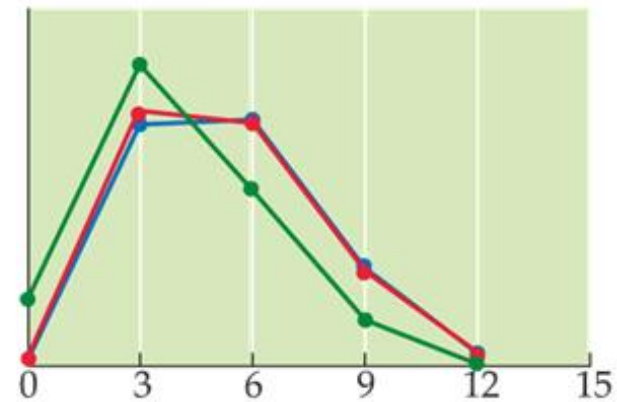
(B)



(C)



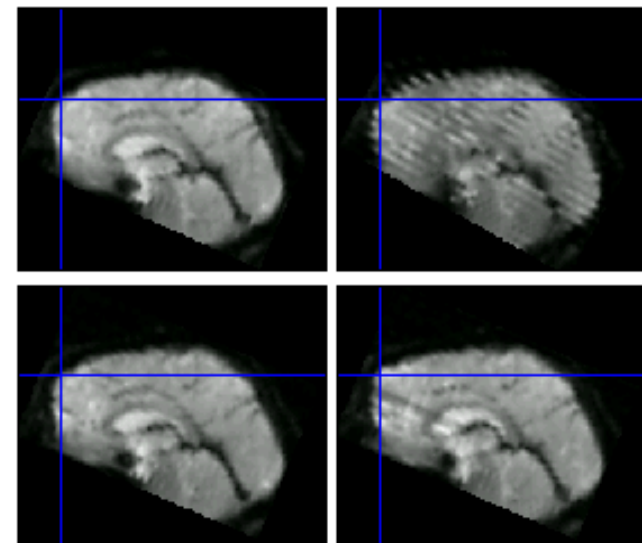
(D)



# SLICE TIMING CORRECTION

Most people now suggest not to do it

- Not all that helpful & requires interpolation
- It may worsen artefacts (e.g., smearing spikes)
- Interacts in unpredictable ways with motion correction
- We spatially smooth across proximal slices
- Mismatching TR and task
- Include temporal derivative of HRF
- What order? Ascending, descending, contiguous, interleaved.



# PREPROCESSING

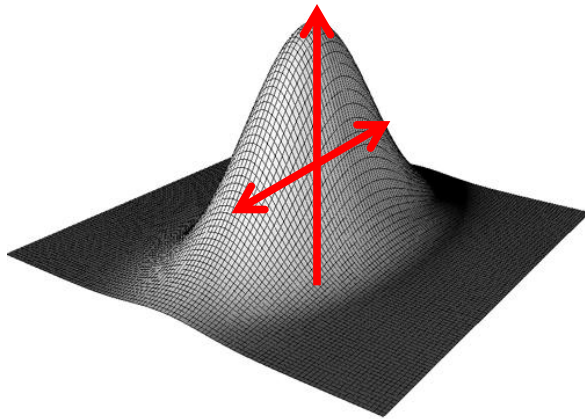
- i. Motion correction
- ii. Slice timing correction
- iii. Spatial filtering**
- iv. Temporal filtering
- v. Intensity normalization



# SPATIAL FILTERING

Replace each voxel's value with a weighted average of its value and the value of its neighbouring voxels.

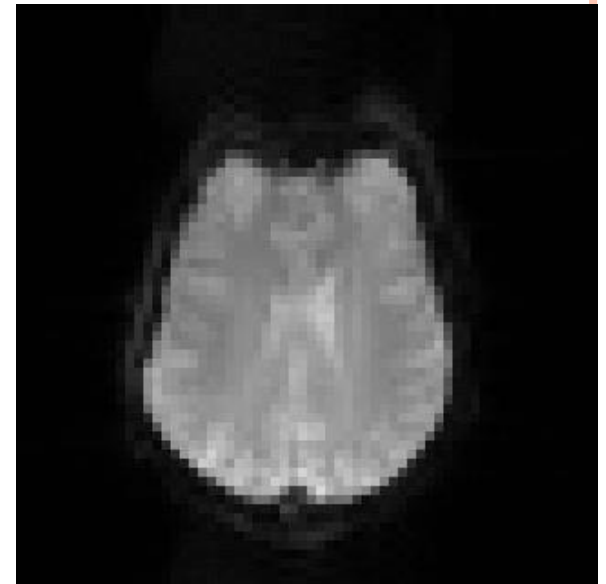
- Gaussian kernel (mm FWHM)



Weights

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 ↔



# 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

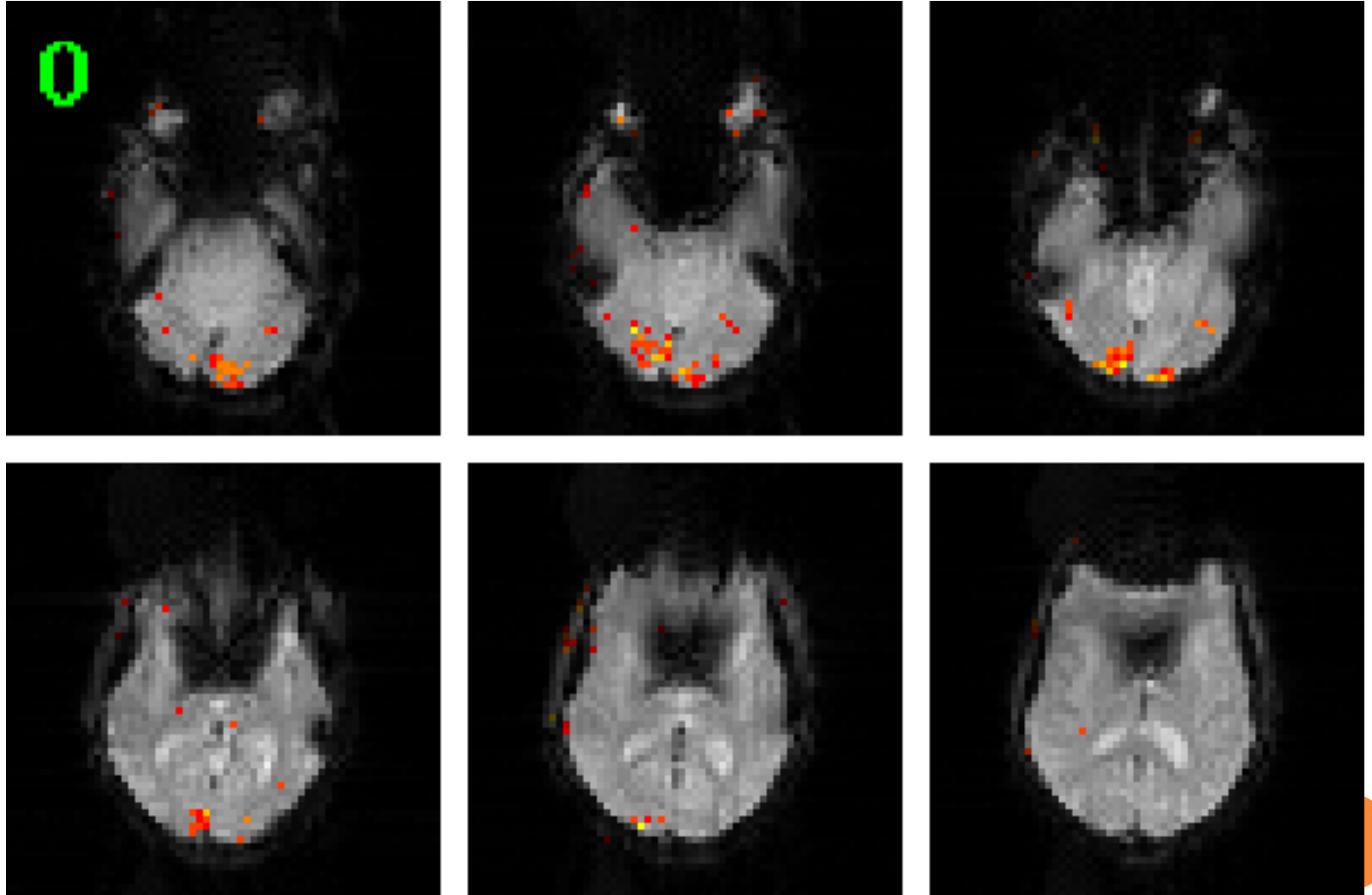
## Disadvantages

- Reduces spatial resolution

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



# SPATIAL FILTERING



Source FSL website

# PREPROCESSING

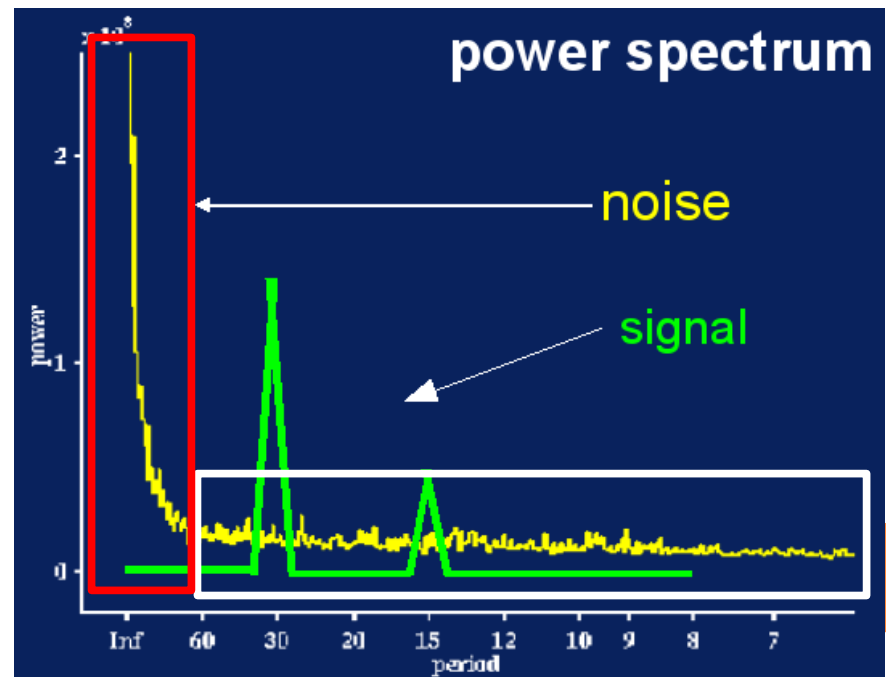
- i. Motion correction
- ii. Slice timing correction
- iii. Spatial filtering
- iv. Temporal filtering**
- v. Intensity normalization



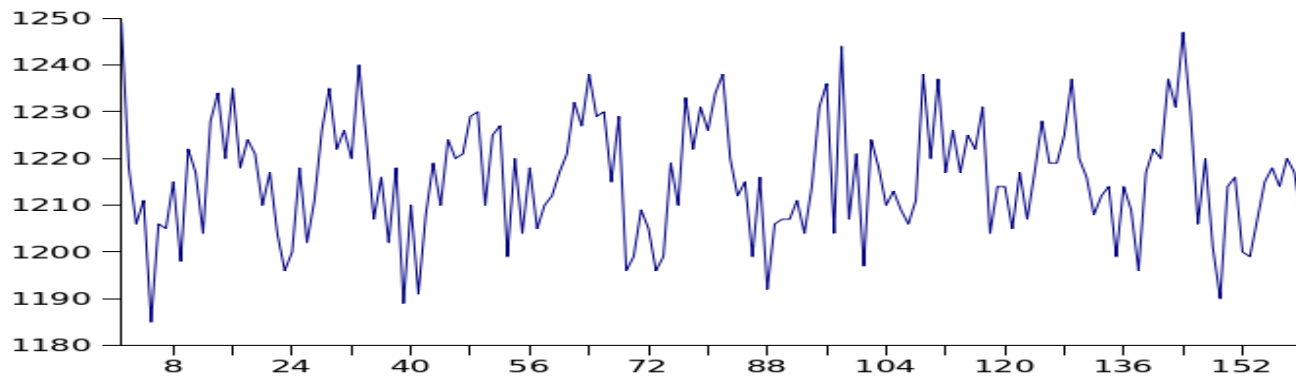
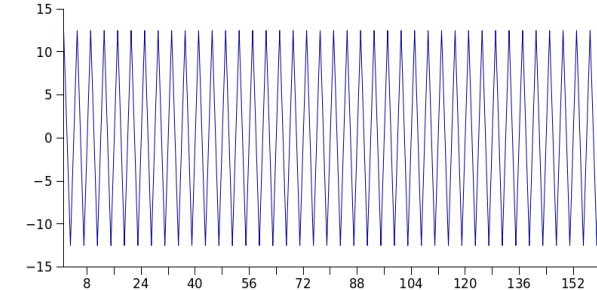
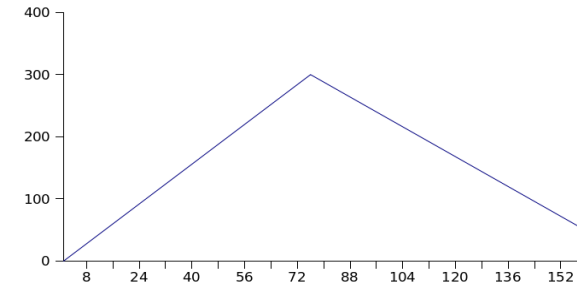
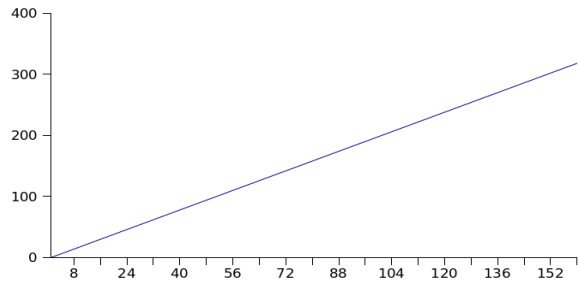
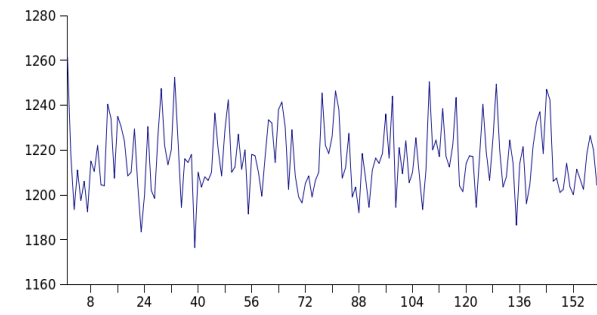
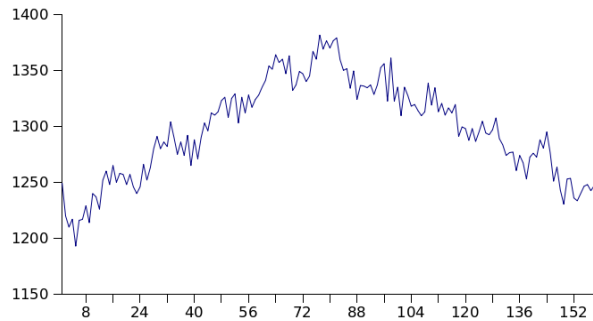
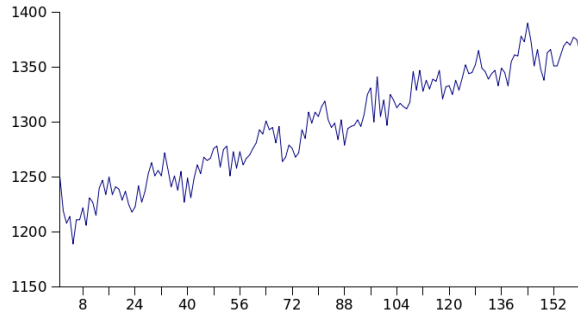


# TEMPORAL FILTERING

- 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:  $1/f$ ):
  - Thermal noise
  - Heart beat
  - Respiration
  - Alertness
  - Learning

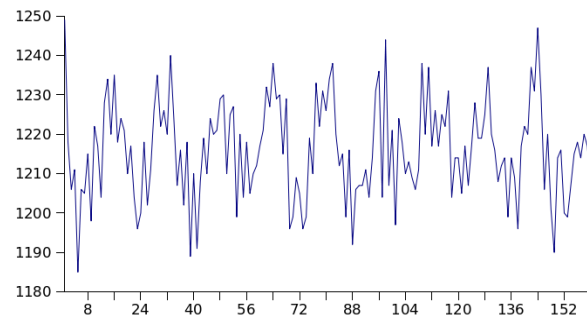
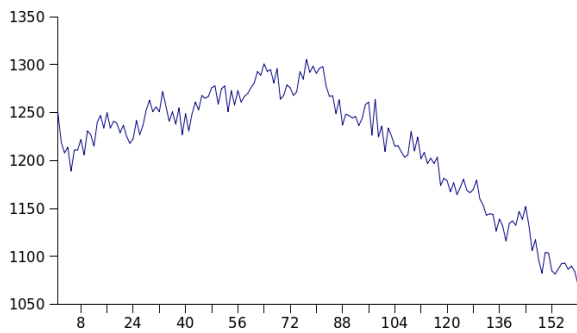


# SIGNAL & NOISE

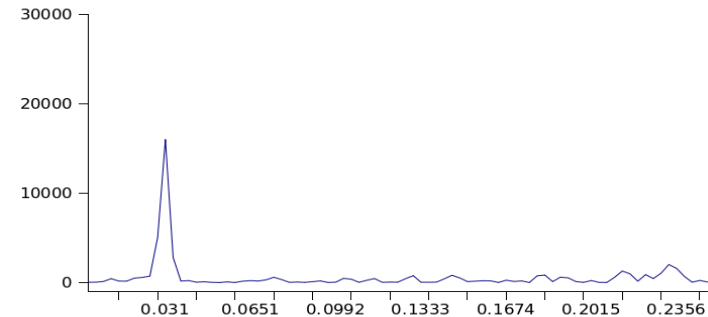
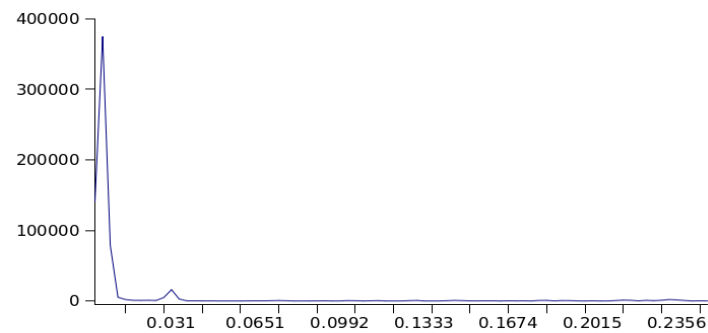


# HIGH-PASS FILTERING

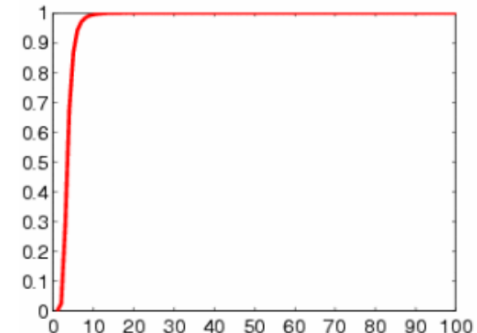
## Timecourse



## Power Spectrum

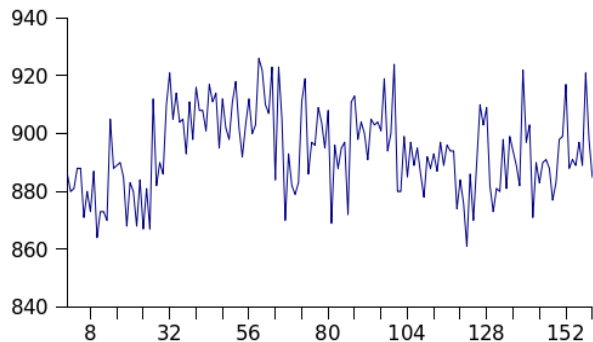
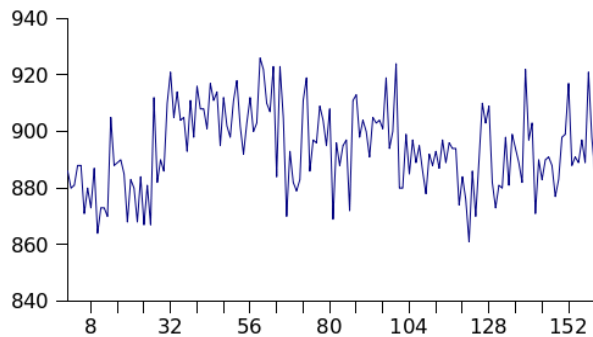


## HP Filter

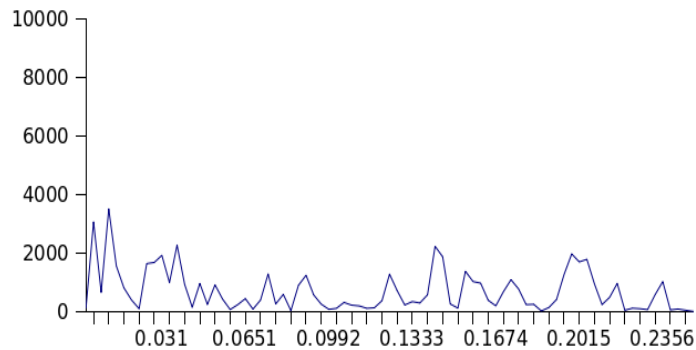
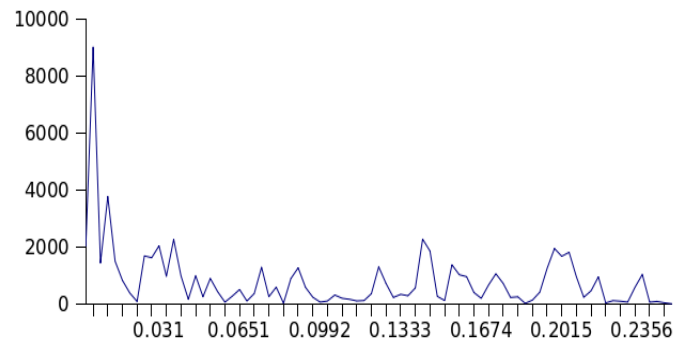


# HIGH-PASS FILTERING

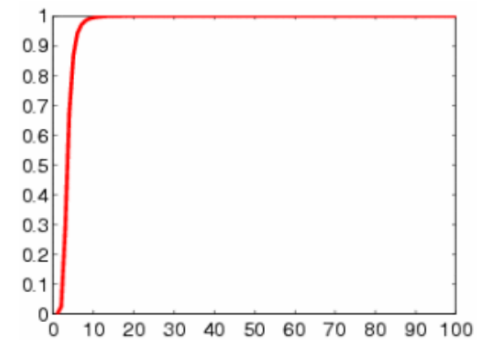
## Timecourse



## Power Spectrum

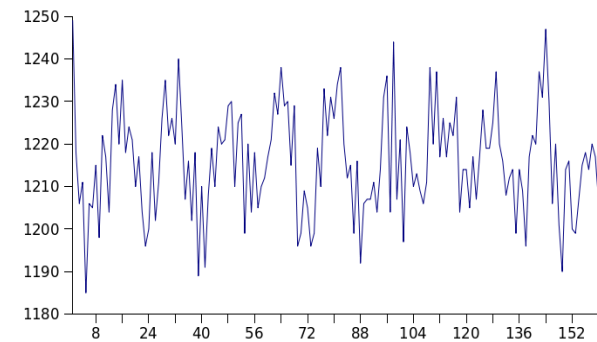
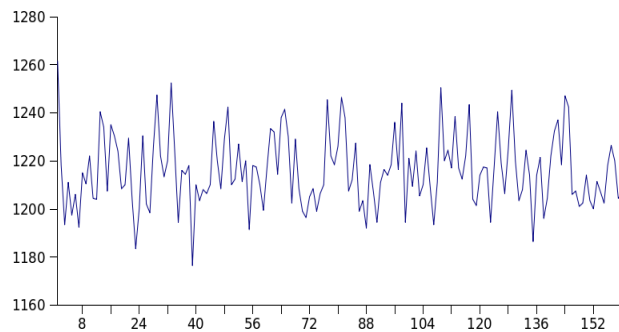


## HP Filter

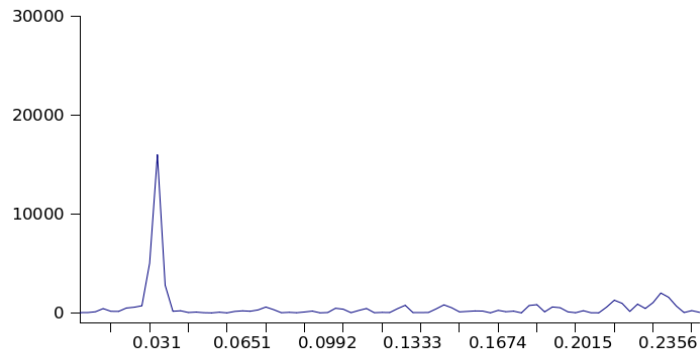
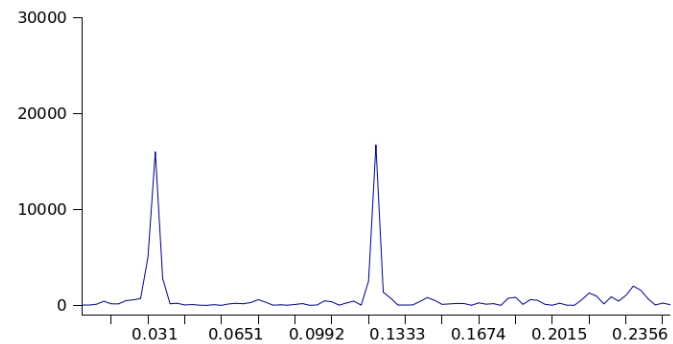


# LOW-PASS FILTERING

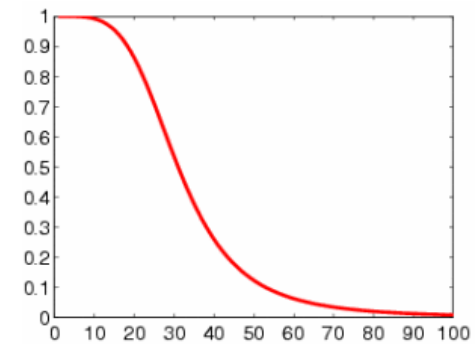
## Timecourse



## Power Spectrum

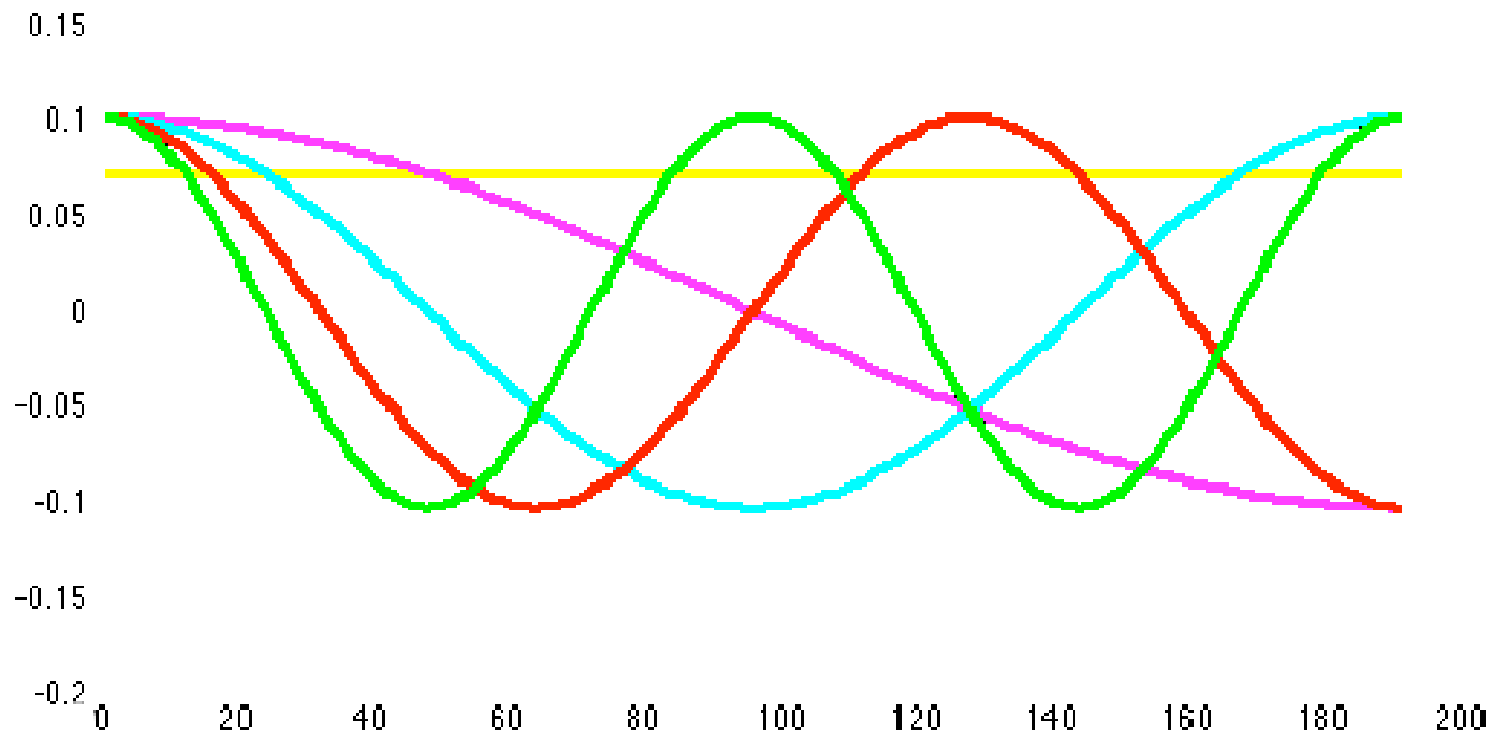


## LP Filter



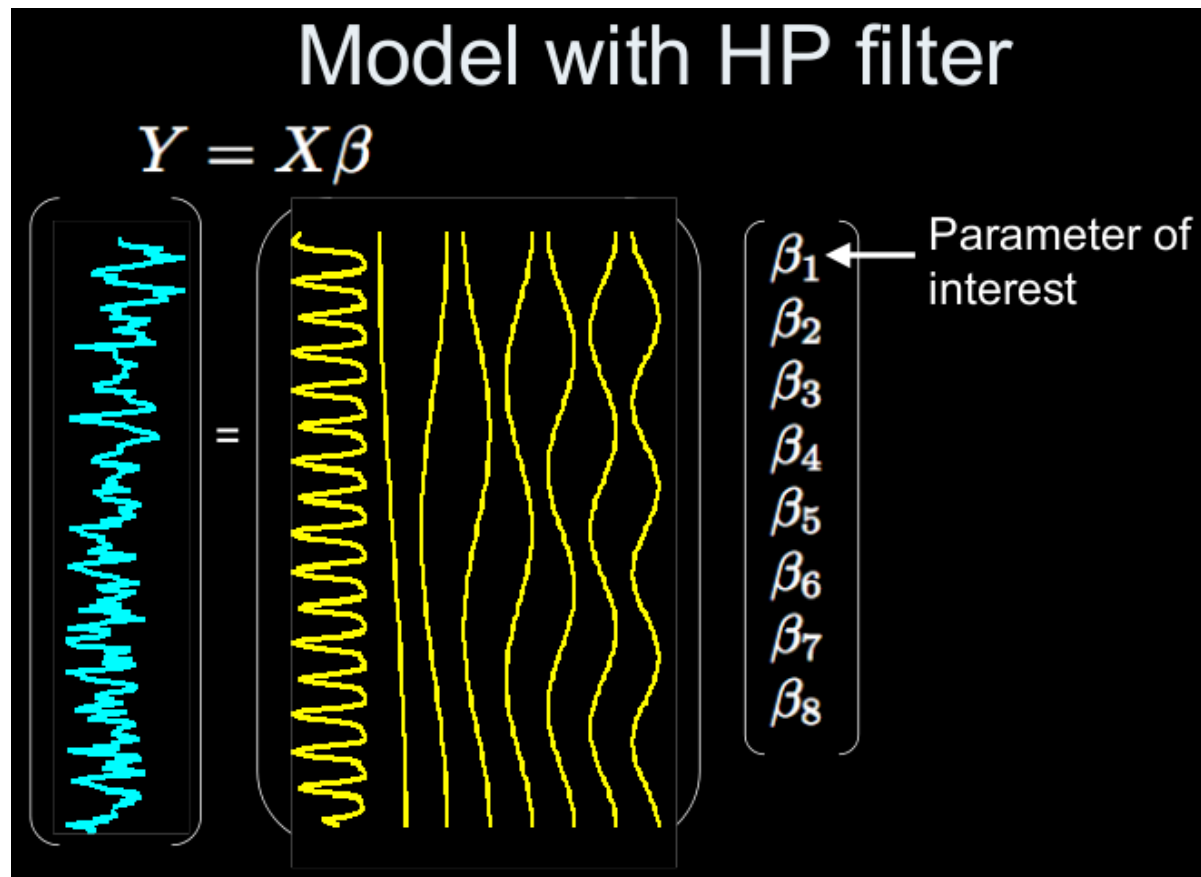
# HP FILTERING STRATEGY I: SPM

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



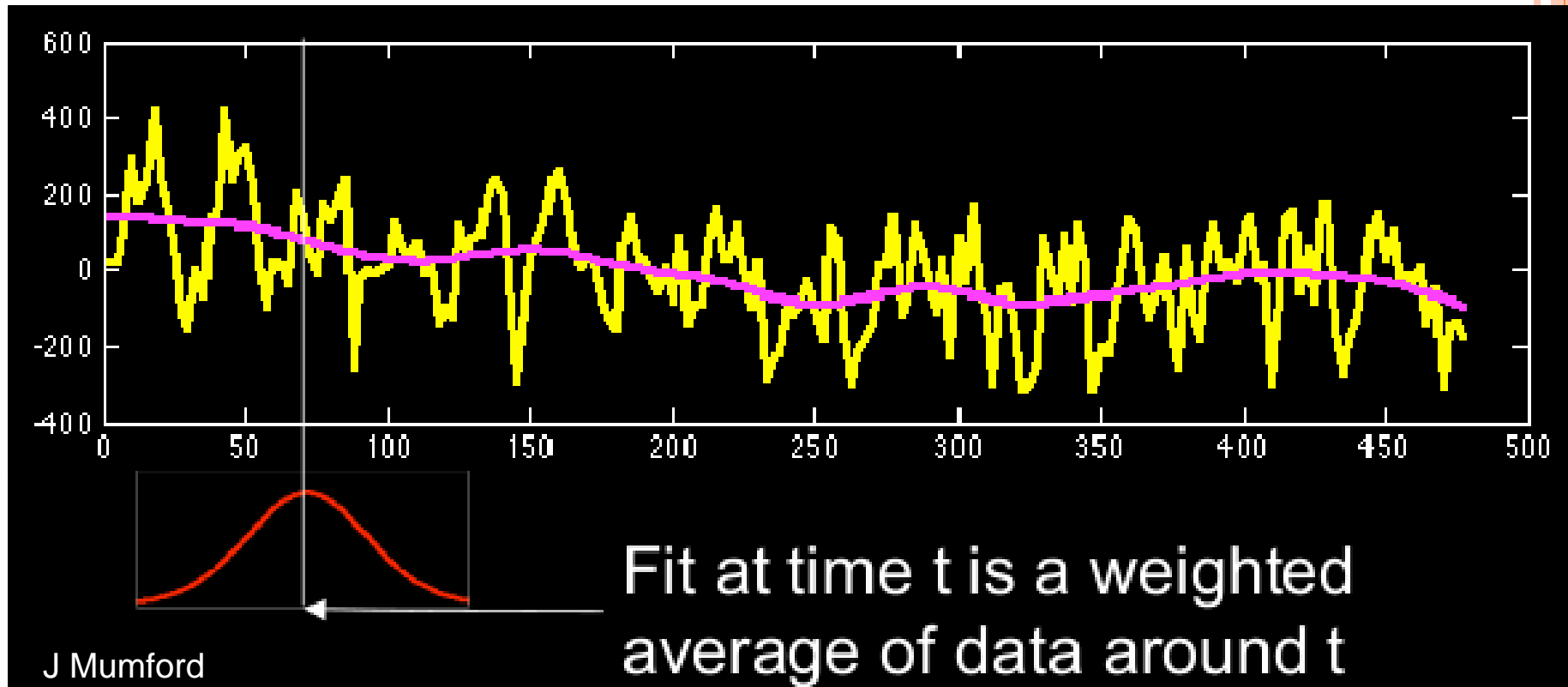
# HP FILTERING STRATEGY I: SPM

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



# HP FILTERING STRATEGY II: FSL

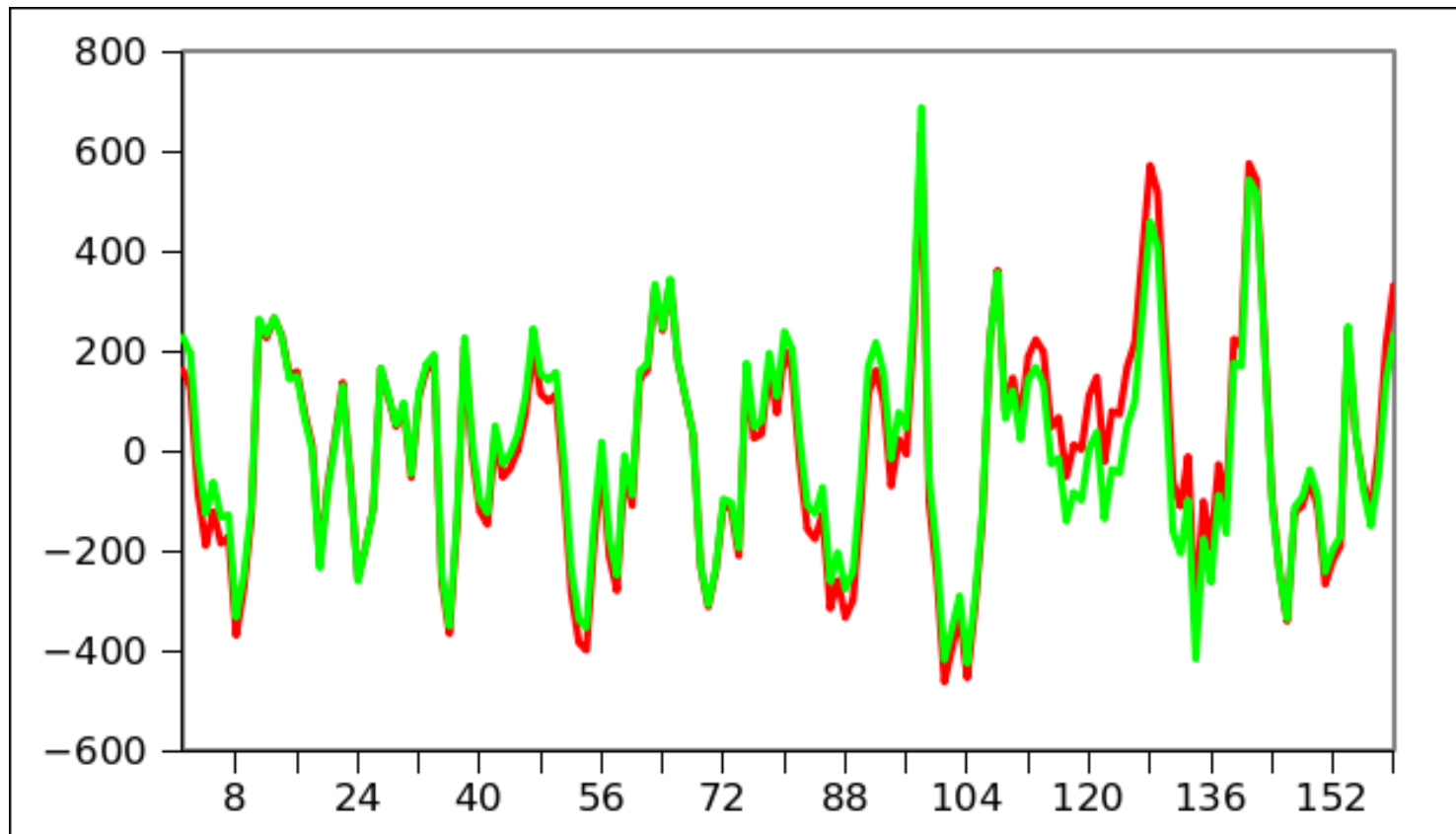
- Remove low drifts from the signal:
  - i. Fit a Gaussian-weighted running line





# HP FILTERING STRATEGY II: FSL

- Remove low drifts from the signal:
  - i. Fit a Gaussian-weighted running line
  - ii. Subtract from data (red is pre-HPF, green is post-HPF)



# PREPROCESSING

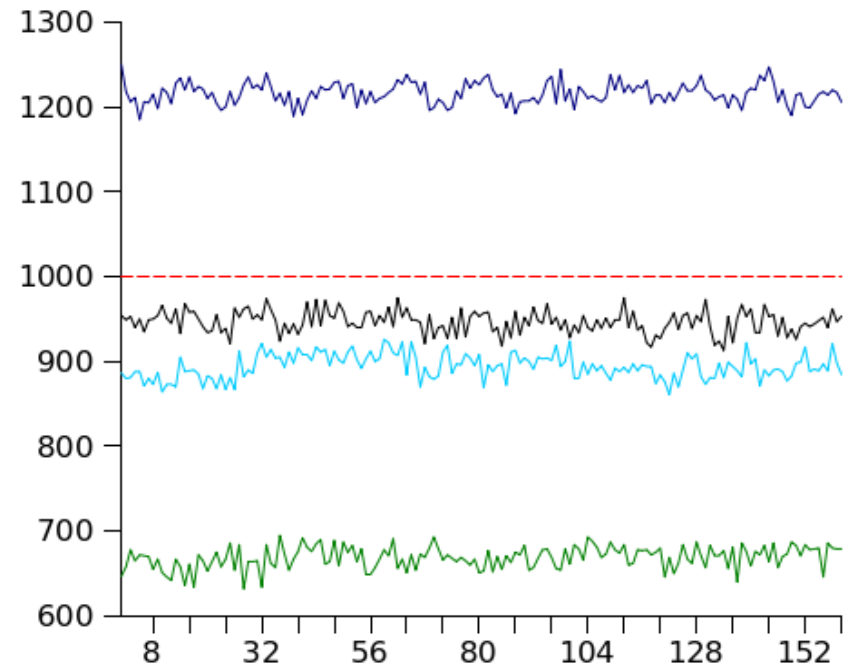
- i. Motion correction
- ii. Slice timing correction
- iii. Spatial filtering
- iv. Temporal filtering
- v. Intensity normalization**



# INTENSITY NORMALIZATION I (GMS)

## *Between-session (grand mean scaling)*

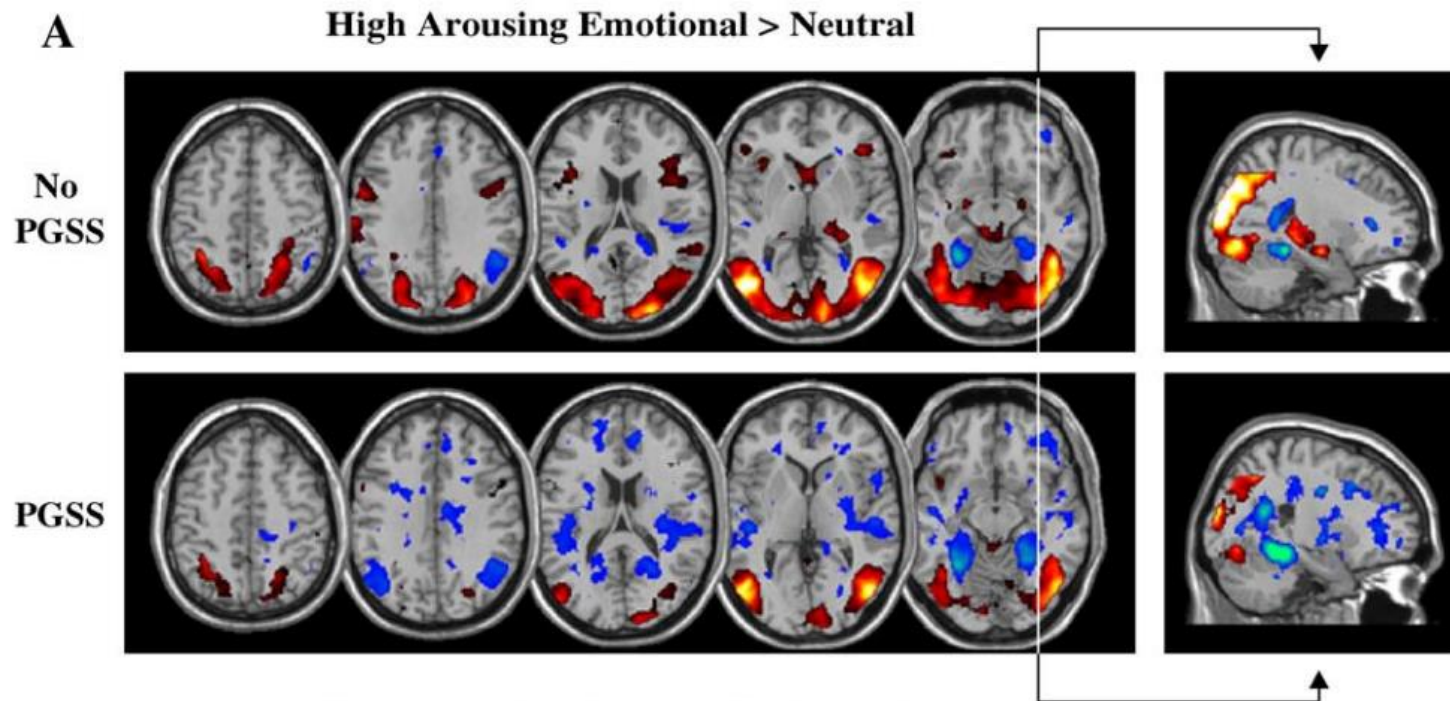
- The mean intensity of each 4D dataset varies for non-experimentally interesting reasons.
- Scale each 4D time-series by a single factor.
- Time-series from different runs are now centred around the same mean.



# INTENSITY NORMALIZATION II

## *Within-session*

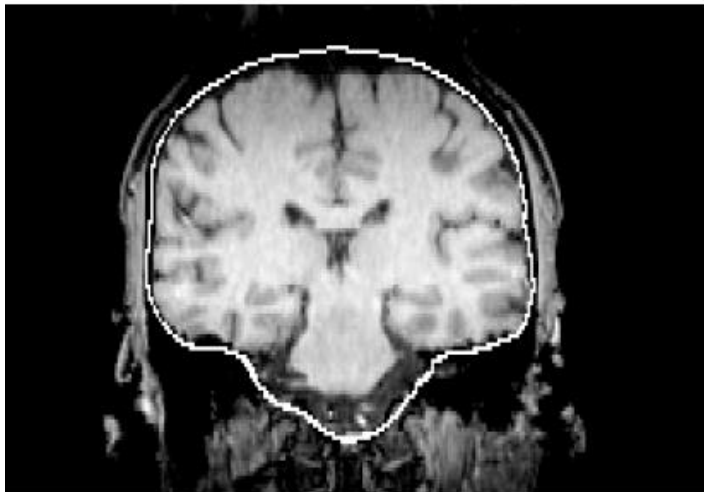
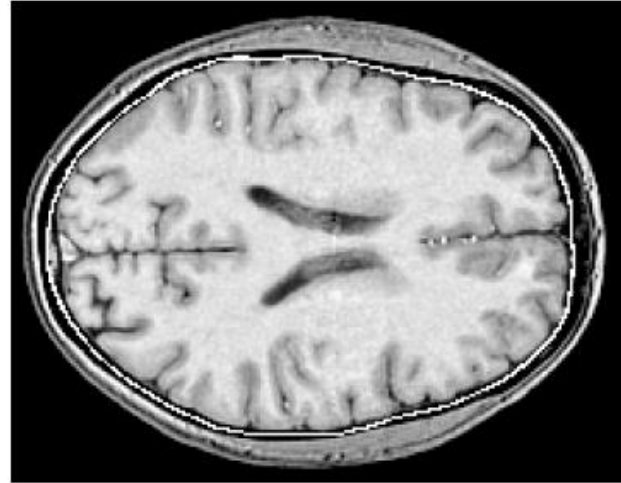
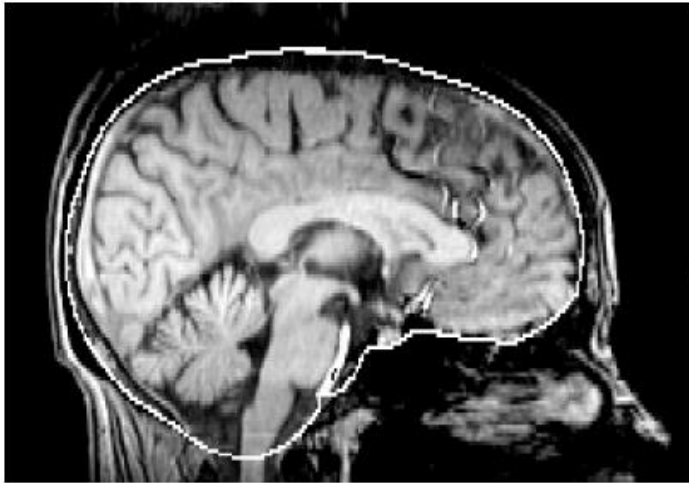
- Forces each volumes (*within a run*) to have the same mean intensity.



Junghofer et al, 2005, neuroImage

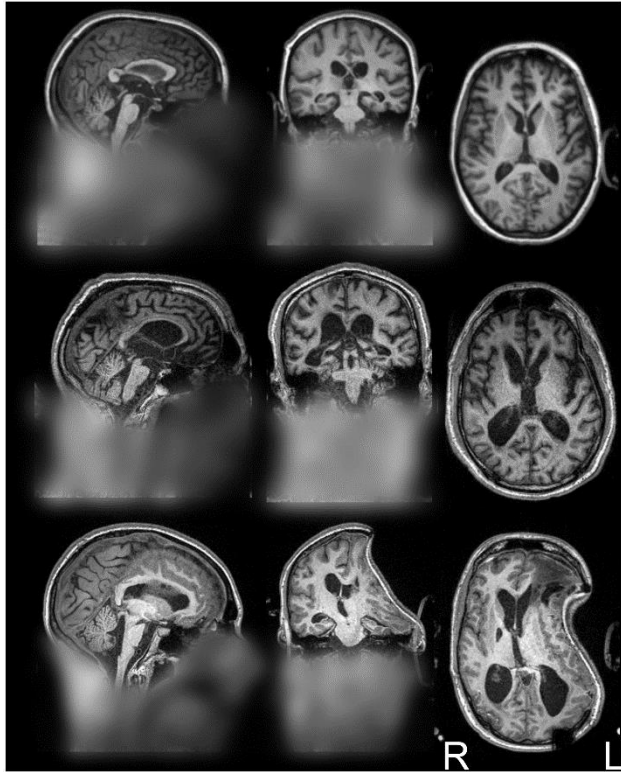


# BRAIN EXTRACTION

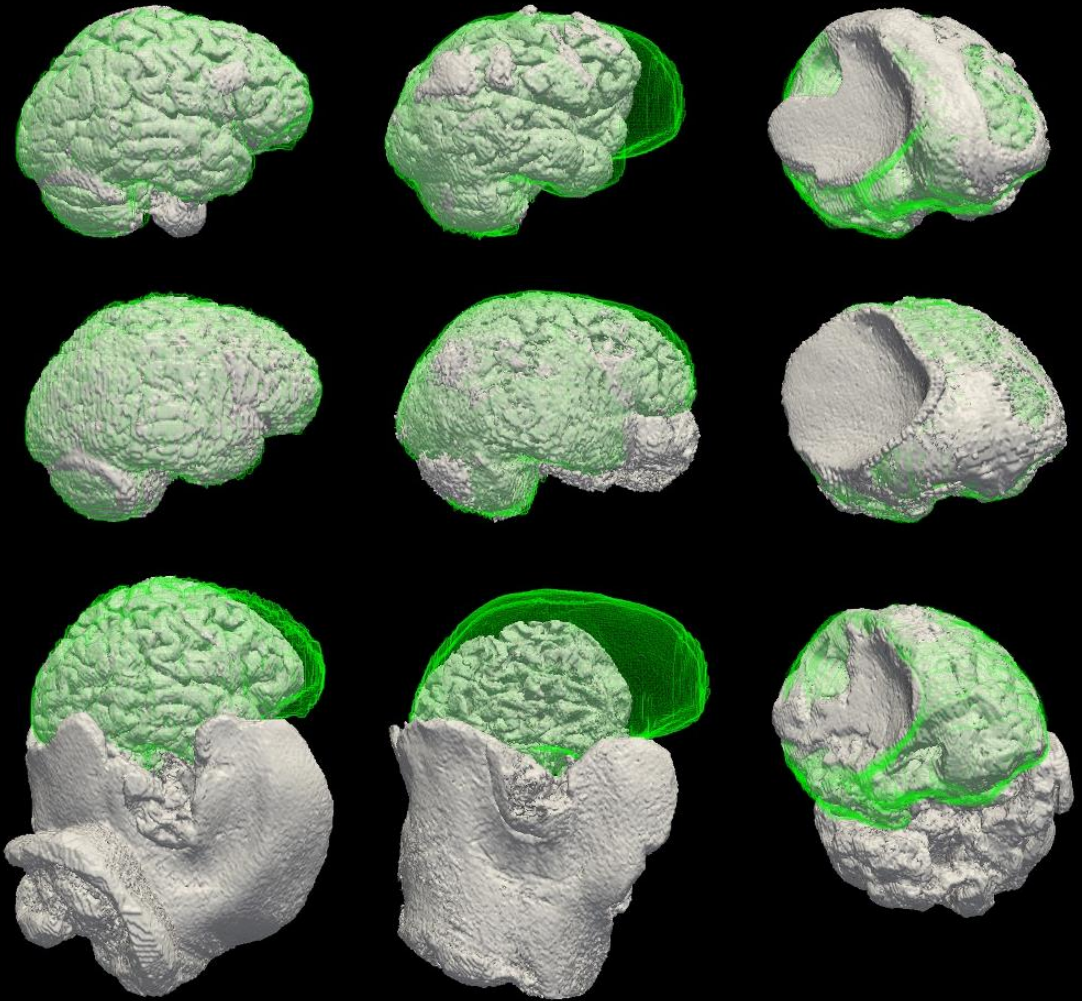




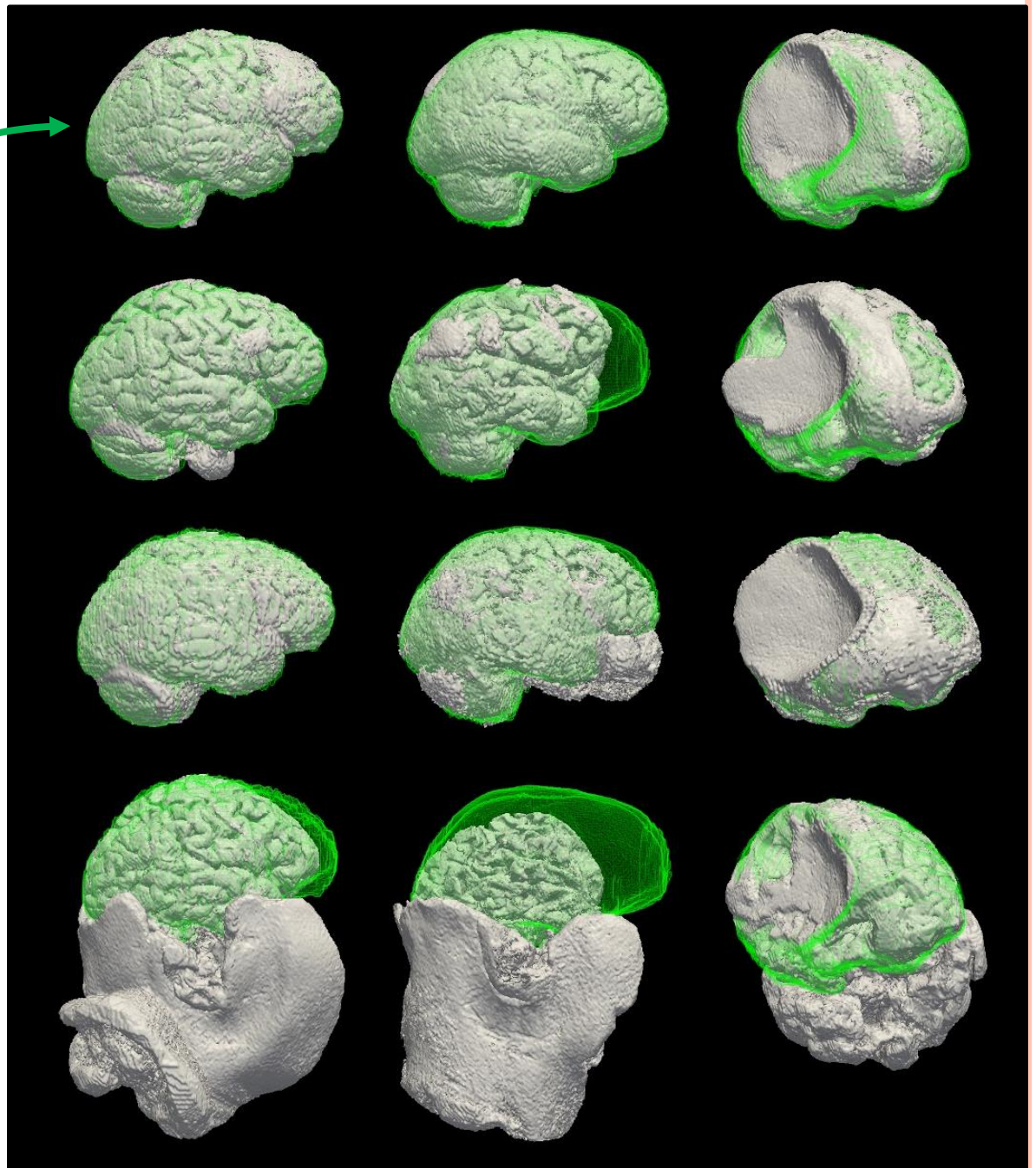
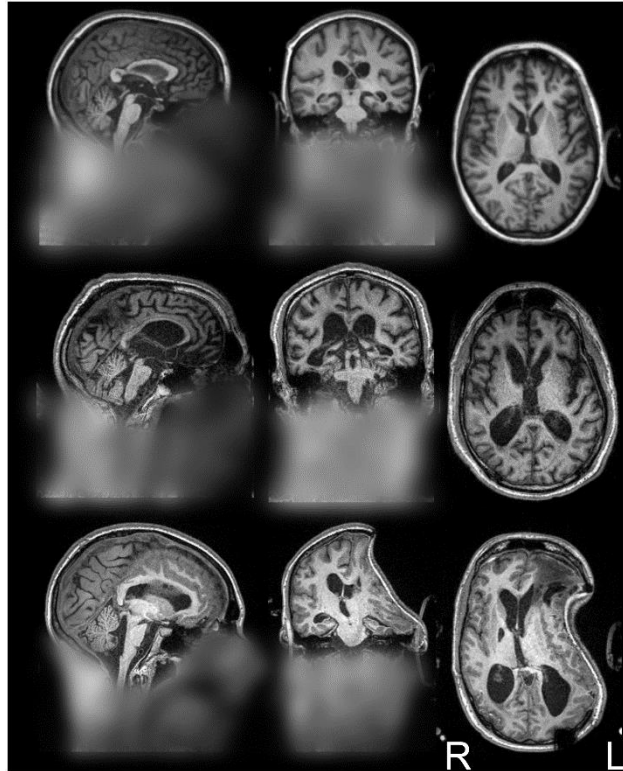
# OPTIBET



## Standard available tools

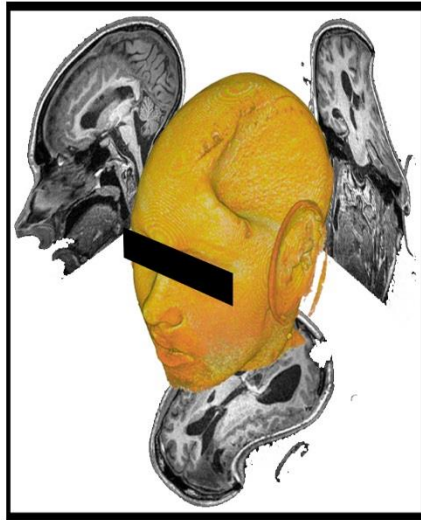


# OPTIBET

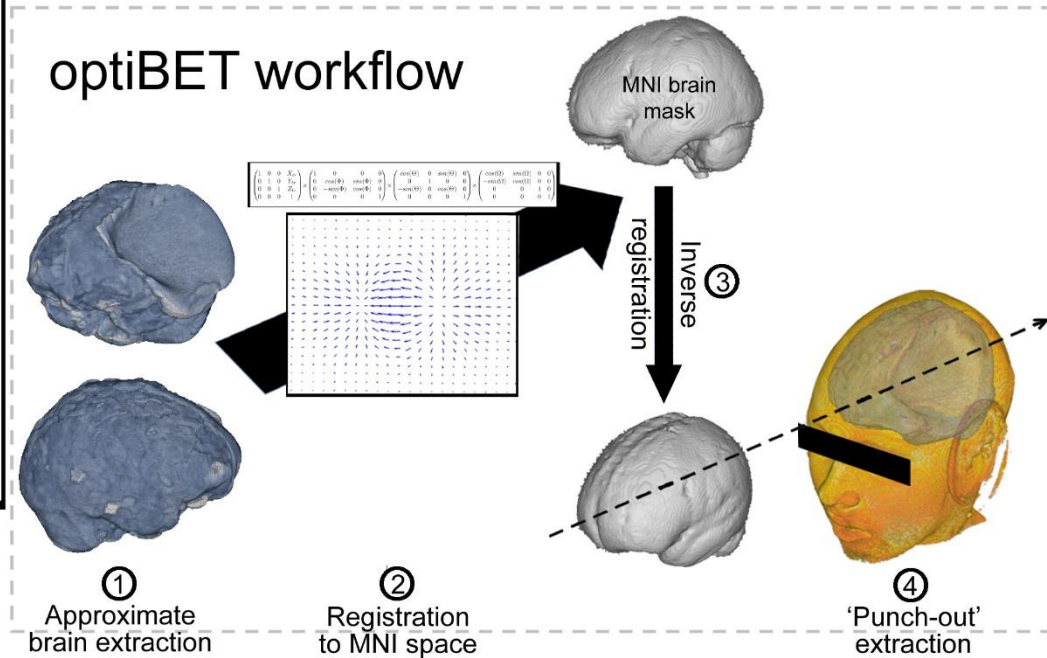




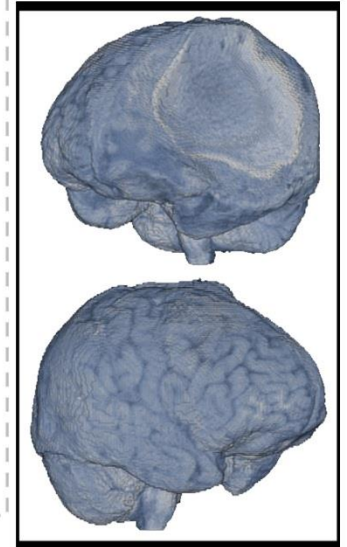
# OPTIBET



Initial head  
image

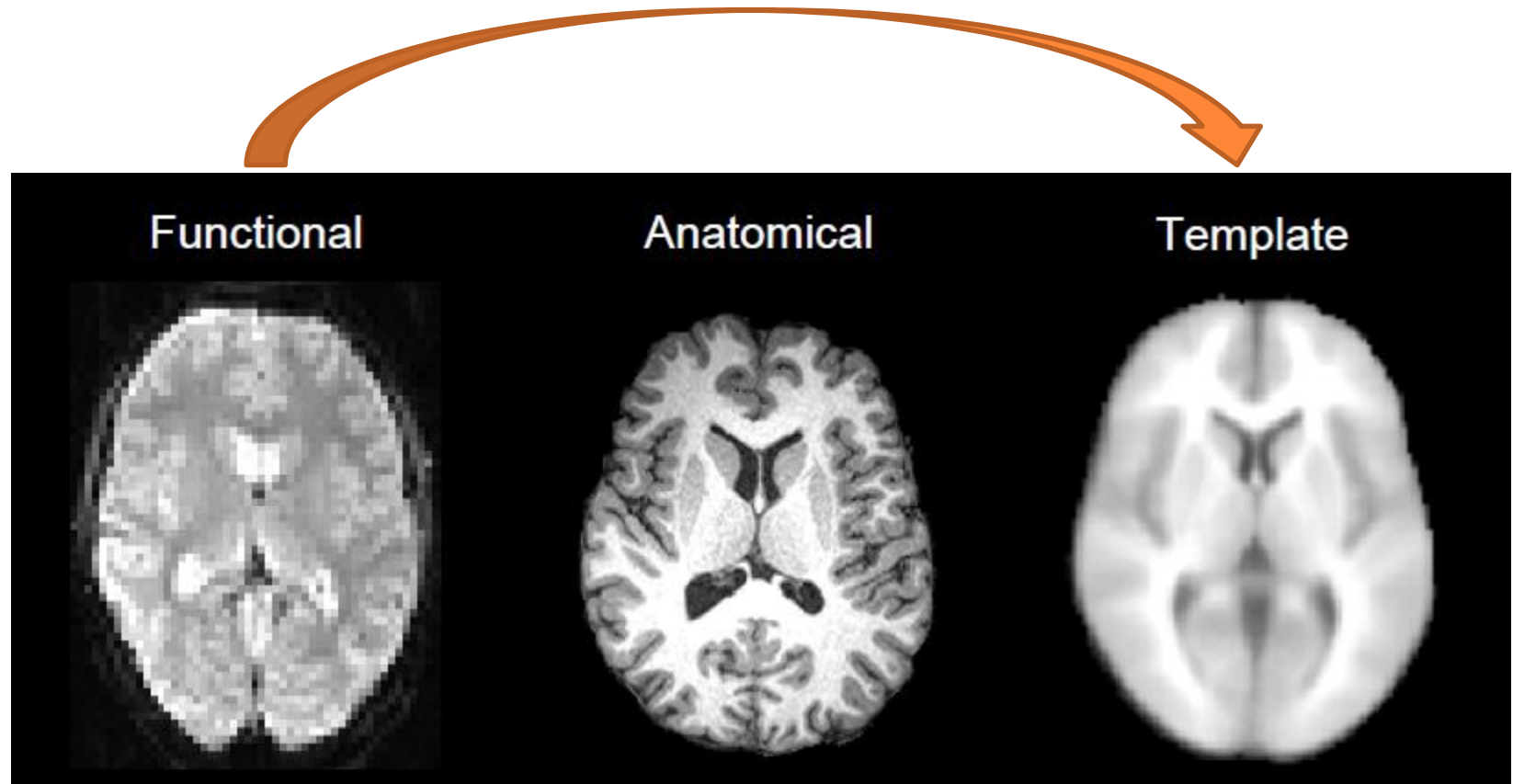


Final brain  
extraction



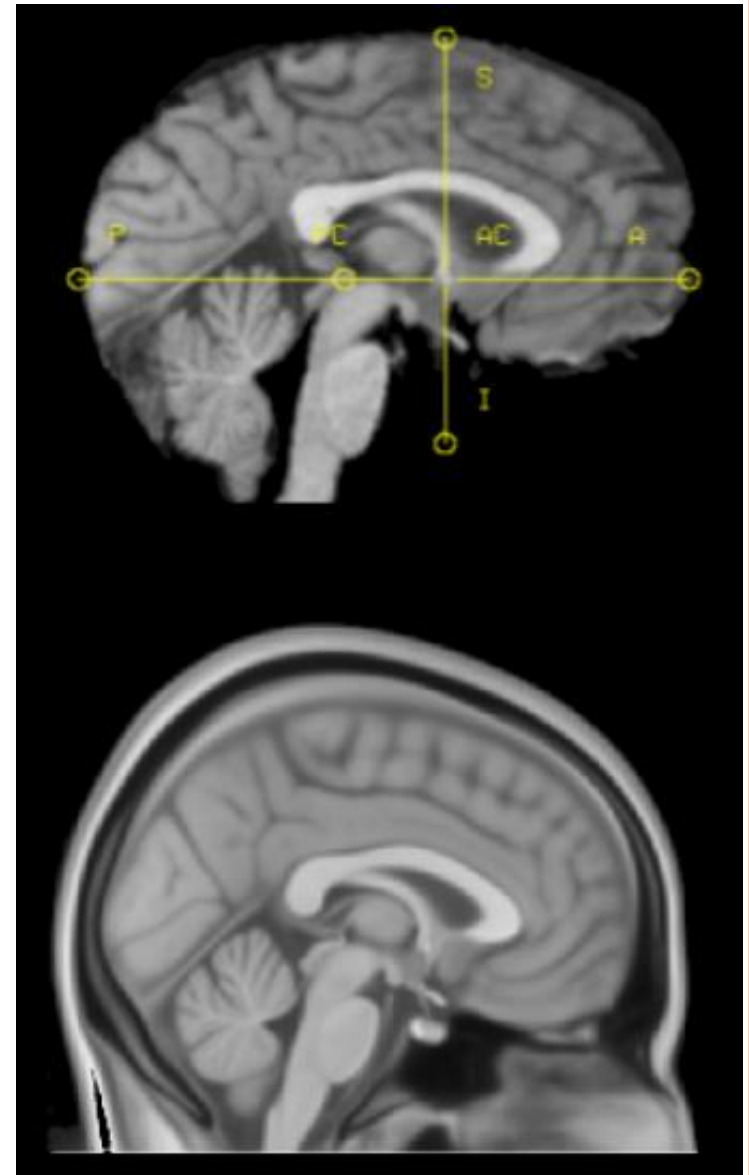


# REGISTRATION

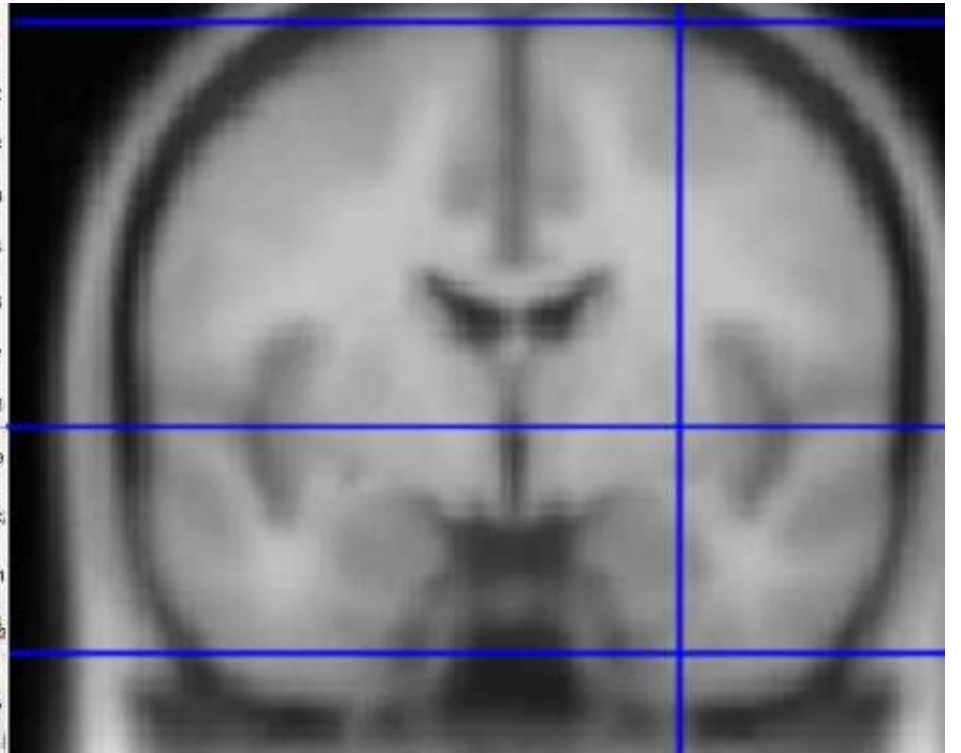
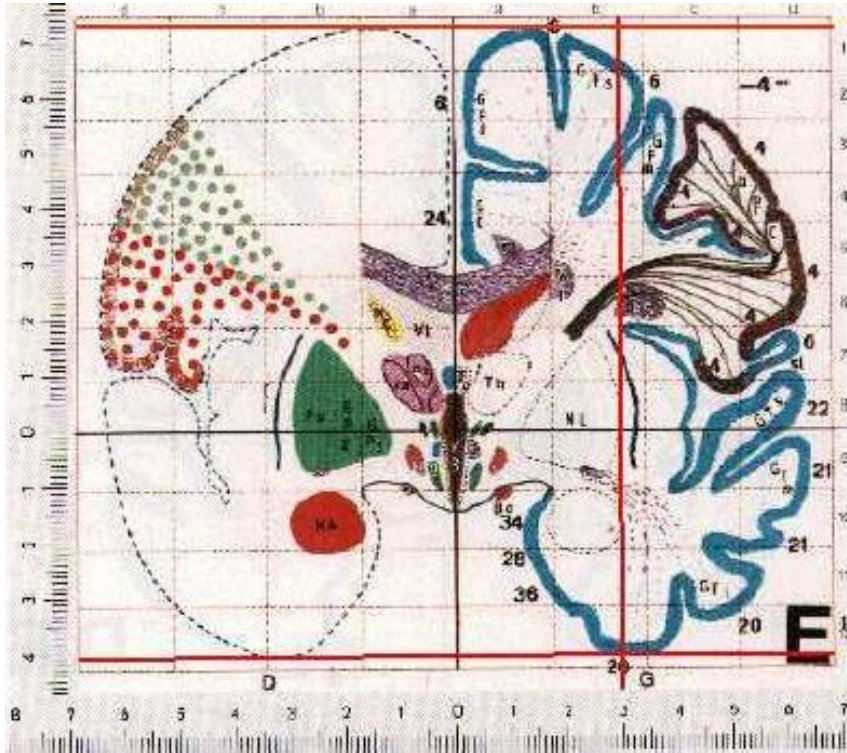


# STANDARD SPACE

- 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”?

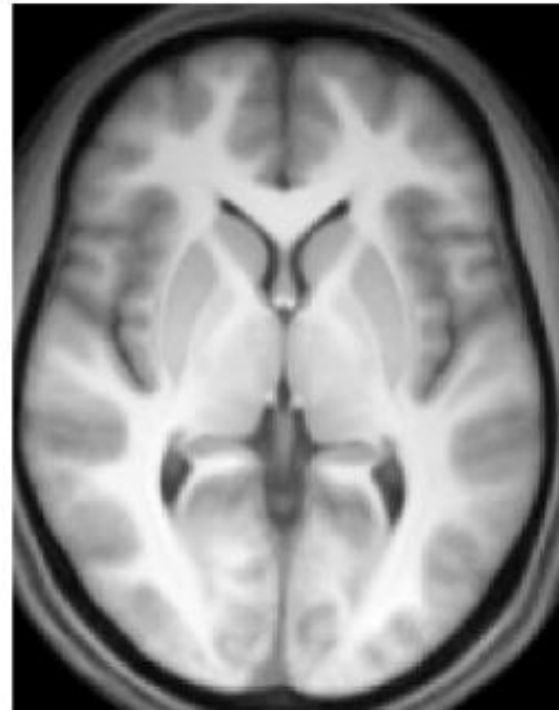
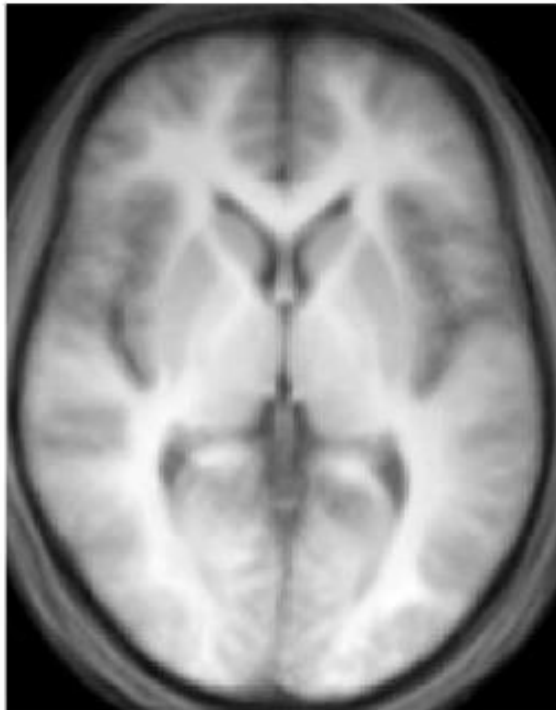


# TRANSFORMATION

- 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 sheers/skews
  - Typically used for registering a subject to the template

# TRANSFORMATION

- Non linear ( $>12\text{dof}$ ):
  - Can be local
  - Can be constrained (e.g., regularization, topology preservation)



# TRANSFORMATION

- 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.





# FLIRT: Cost Functions

Important: Allowable image modalities

Less important: Details

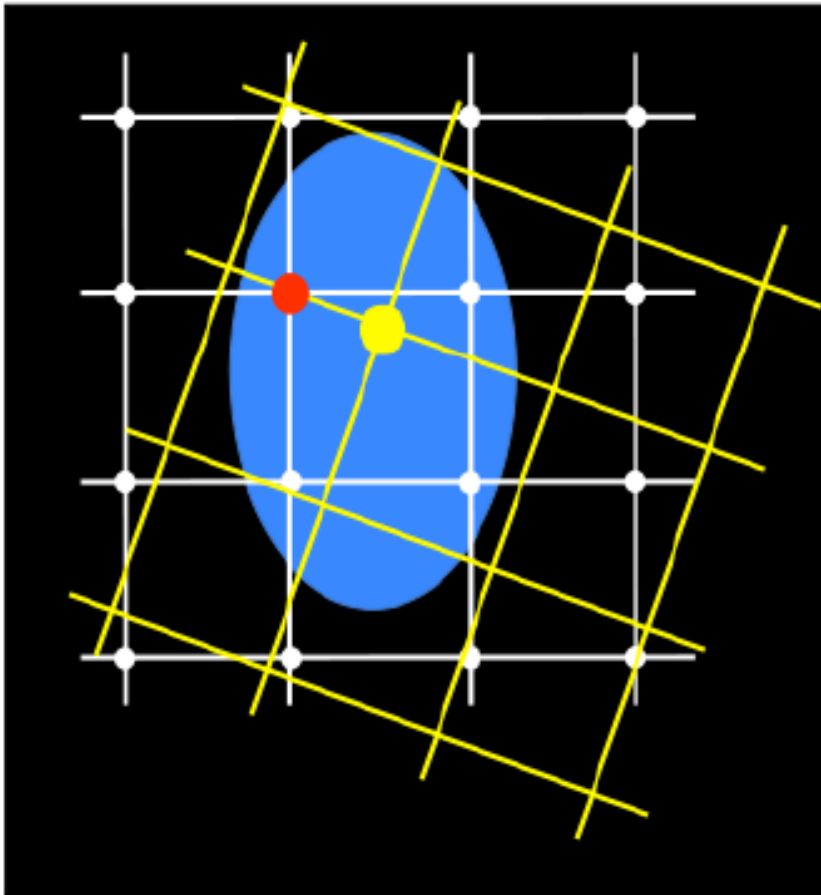
Least Squares	<b><i>Same modality</i></b> (exact sequence parameters)
Normalised Correlation	<b><i>Same modality</i></b> (can change brightness & contrast)
Correlation Ratio	<b><i>Any MR modalities</i></b>
Mutual Information	<b><i>Any modalities</i></b> (including CT, PET, etc.)
Normalised Mutual Info.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





# Interpolation

Finds intensity values between grid points



Various types include

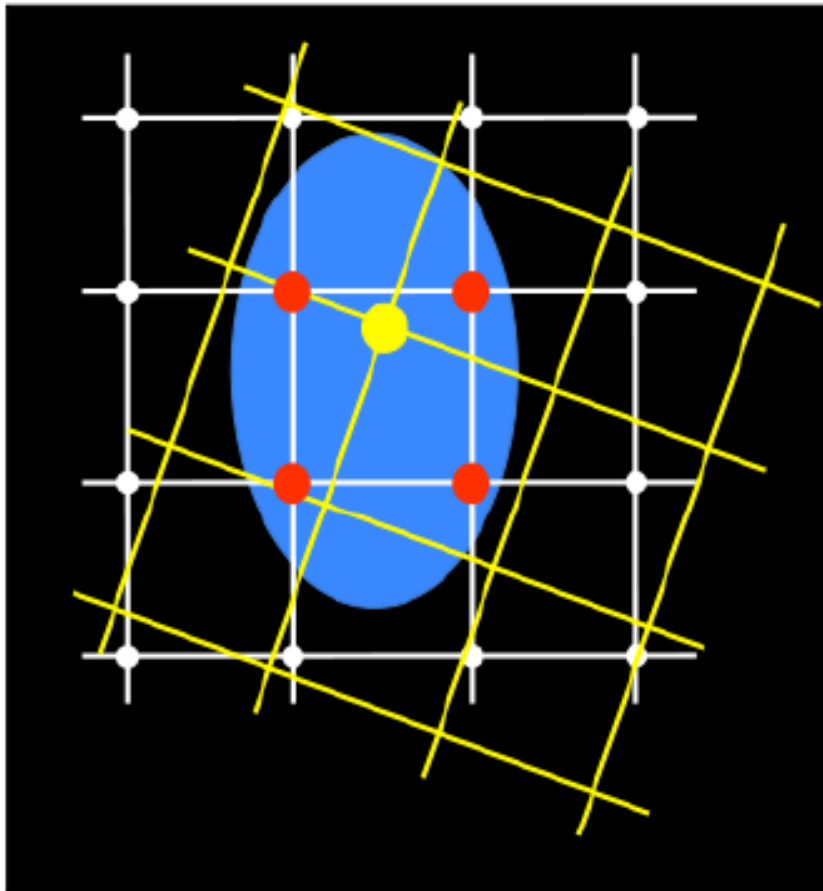
- Nearest Neighbour
- Trilinear
- Sinc
- Spline
- k-Space methods





# Interpolation

Finds intensity values between grid points



Various types include

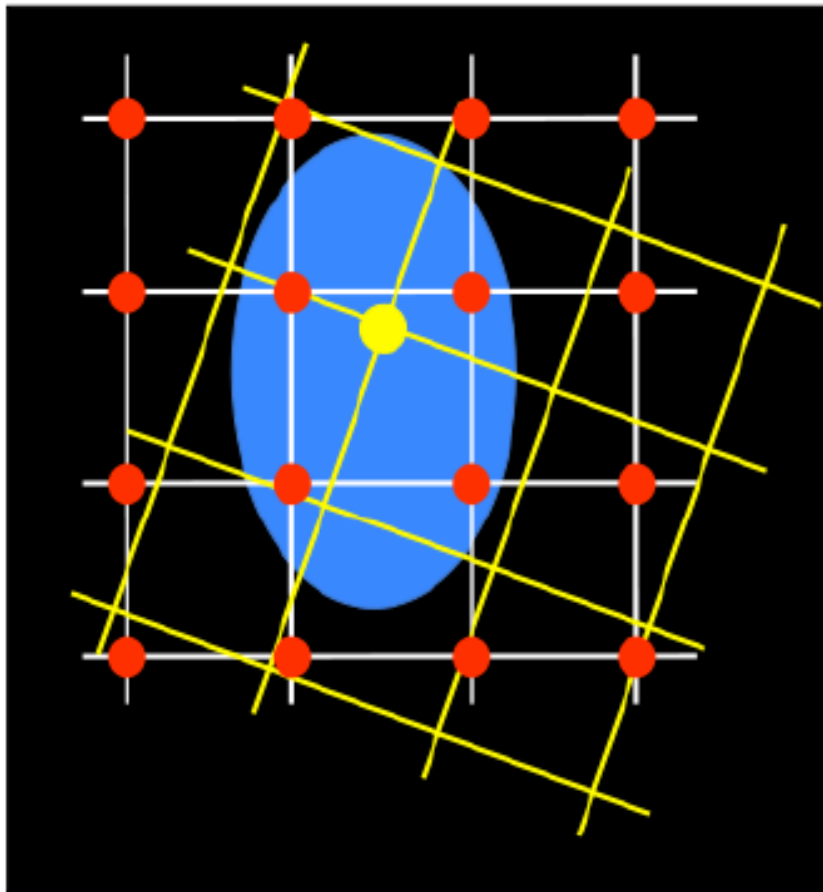
- Nearest Neighbour
- **Trilinear**
- Sinc
- Spline
- k-Space methods





# Interpolation

Finds intensity values between grid points



Various types include

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

Considerations: speed, accuracy, stability

