

# fMRI Preprocessing

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#### Contents

#### \* Preliminaries

- \* Rigid-body and affine transformations
- Optimisation and objective functions
- \* Transformations and interpolation
- \* Within-subject: Realignment
- \* Within-subject: EPI Distortion Correction
- \* Within-subject: Coregistration
- \* Between-subject: Normalise/Segment
- \* Between-subject: Dartel
- \* Between-subject: Smoothing

#### Image registration

Most "preprocessing" involves aligning images together.

#### Two components:

- Registration i.e. Optimise the parameters that describe spatial transformations between the images.
- **Transformation** i.e. Re-sample according to the determined transformation parameters.

## **Rigid-body transformations**

- \* Assume that brain of the same subject doesn't change shape or size in the scanner.
  - \* Head can move, but remains the same shape and size.
  - \* Some exceptions:
    - \* Image distortions.
    - \* Brain slops about slightly because of gravity.
    - \* Brain growth or atrophy over time.
- \* If the subject's head moves, we need to correct the images.
  - \* Do this by image registration.

### 2D affine transforms

\* Translations by t<sub>x</sub> and t<sub>y</sub>

$$x_1 = x_0 + t_x$$

\* 
$$y_1 = y_0 + t_y$$

\* Rotation around the origin by  $\Theta$  radians  $\overline{\Theta}$ 

\*Shear

 $x_1 = x_0 + h y_0$ 

 $= y_0$ 

$$x_1 = \cos(\Theta) x_0 + \sin(\Theta) y_0$$

\* 
$$y_1 = -\sin(\Theta) x_0 + \cos(\Theta) y_0$$

- \* Zooms by  $s_x$  and  $s_y$ 
  - \*  $x_1 = s_x x_0$ \*  $y_1 = s_y y_0$



### 2D affine transforms

\* Translations by t<sub>x</sub> and t<sub>y</sub>

\* 
$$x_1 = 1 x_0 + 0 y_0 + t_x$$

\* 
$$y_1 = 0 x_0 + 1 y_0 + t_y$$

\* Rotation around the origin by  $\Theta$  radians  $\overline{\Theta}$ 

\*Shear

 $x_1 = 1 x_0 + h y_0 + 0$ 

 $*y_1 = 0 x_0 + 1 y_0 + 0$ 

$$x_1 = \cos(\Theta) x_0 + \sin(\Theta) y_0 + 0$$

$$y_1 = -\sin(\Theta) x_0 + \cos(\Theta) y_0 + C$$

- \* Zooms by  $s_x$  and  $s_y$ :
  - \*  $x_1 = s_x x_0 + 0 y_0 + 0$
  - \*  $y_1 = 0 x_0 + s_y y_0 + 0$



#### 3D rigid-body transformations

- \* A 3D rigid body transform is defined by:
  - \* 3 translations in X, Y & Z directions
  - \* 3 rotations about X, Y & Z axes
- \* The order of the operations matters



#### Voxel-to-world transforms

- \* Affine transform associated with each image
  - \* Maps from voxels (x=1..n<sub>x</sub>, y=1..n<sub>y</sub>, z=1..n<sub>z</sub>) to some world coordinate system. e.g.,
    - \* Scanner co-ordinates images from DICOM toolbox
    - \* T&T/MNI coordinates spatially normalised
- \* Registering image B (source) to image A (target) will update B's voxel-to-world mapping
  - \* Mapping from voxels in A to voxels in B is by
    - \* A-to-world using  $M_A$ , then world-to-B using  $M_B^{-1}$
    - \* M<sub>B</sub><sup>-1</sup> M<sub>A</sub>

#### Optimisation

- \* Image registration is done by optimisation.
- Optimisation involves finding some "best" parameters according to an "objective function", which is either minimised or maximised
- The "objective function" is often related to a probability based on some model



# Optimisation

\* Because registration only finds a *local optimum*, some manual reorienting of the images may be needed before doing anything else in SPM.

> An MNI-space image from spm12/canonical directory.



#### **Objective functions**



#### \* Intra-modal

- \* Mean squared difference (minimise)
- \* Normalised cross correlation (maximise)



- \* Inter-modal (or intra-modal)
  - \* Mutual information (maximise)
  - \* Normalised mutual information (maximise)
  - \* Entropy correlation coefficient (maximise)

## Simple interpolation

- \* Nearest neighbour
  - Take the value of the closest voxel
- \* Tri-linear
  - \* Just a weighted average of the neighbouring voxels
  - \*  $f_5 = f_1 x_2 + f_2 x_1$ \*  $f_6 = f_3 x_2 + f_4 x_1$ \*  $f_7 = f_5 y_2 + f_6 y_1$



#### **B-spline interpolation**





#### Contents

\* Preliminaries

#### \* Within-subject: Realignment

- Realignment by minimising mean-squared difference
- \* Residual artifacts
- \* Within-subject: Realignment
- \* Within-subject: EPI Distortion Correction
- \* Within-subject: Coregistration
- \* Between-subject: Normalise/Segment
- \* Between-subject: Dartel
- \* Between-subject: Smoothing

#### Mean-squared difference



Original Joint Histogram

Final Joint Histogram

- Minimising mean-squared difference works for intra-modal registration (realignment)
- \* Simple relationship between intensities in one image, versus those in the other
  - \* Assumes normally distributed differences



#### **Motion estimates**









"Browse" option from Check Reg allows fMRI time series to be shown as a movie.

/home/john/fmri/fM00223/fM00223\_074.img,1

▶ > Q

File Edit View Insert Tools Desktop Window SPM Figure Help

#### Residual errors from aligned fMRI

- \* Re-sampling can introduce interpolation errors
  - \* especially tri-linear interpolation
- \* Gaps between slices can cause aliasing artefacts
- \* Slices are not acquired simultaneously
  - \* rapid movements not accounted for by rigid body model
- \* Image artefacts may not move according to a rigid body model
  - image distortion
  - \* image dropout
  - \* Nyquist ghost
- \* BOLD signal changes influence the estimated motion.
- \* Functions of the estimated motion parameters can be modelled as confounds in subsequent analyses

#### Contents

- \* Preliminaries
- \* Within-subject: Realignment

#### \* Within-subject: EPI Distortion Correction

- \* FieldMap Toolbox
- \* Movement by distortion interaction
- \* Within-subject: Coregistration
- \* Between-subject: Normalise/Segment
- \* Between-subject: Dartel
- \* Between-subject: Smoothing

#### **EPI** distortion

- \*Magnetic susceptibility differs among tissues.
- \*Greatest difference is between air and tissue.
- \*Subject disrupts B<sub>0</sub> field, rendering it inhomogeneous
- \* Distortions in phase-encode direction













#### FieldMap toolbox

- Computes a voxeldisplacement map (VDM) from fieldmap scans.
- Used to correct distortions in EPI.

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Generate unwrapped field maps which are converted to voxel displacement maps (VDM) that can be used to unwarp geometrically distorted EPI images.

The resulting VDM files are saved with the prefix vdm and can be applied to images using Apply VDM or in combination with Realign & Unwarp to calculate and correct for the combined effects of static and movement-related susceptibility induced distortions.

This branch contains 1 items: \* Data

# Phase unwrapping

- \* Phase of complex data used.
- π/2 < phase < π/2</li>
- \* Phase-unwrapping needed.
  - \* Part that is most likely to go wrong.
  - \* Phase is poorly defined when magnitude is small relative to noise.





#### **Movement-by-distortion interaction**

#### Original position



Original position



After rotation



After rotation



# Realign & Unwarp



#### Estimation of EPI deformation fields



Derivative w.r.t. pitch



Derivative w.r.t. roll





## **Correcting for distortion changes**



Estimate movement parameters.



Estimate reference from mean of all scans.

Estimate new distortion fields for each image:

 estimate rate of change of field with respect to the current estimate of movement parameters in pitch and roll.

 $\Delta \varphi + \Delta \theta$  $\partial B_0 / \partial \varphi \partial B_0 / \partial \theta$ 



Unwarp time series.



#### Andersson et al, 2001

#### Contents

- \* Preliminaries
- \* Within-subject: Realignment
- \* Within-subject: EPI Distortion Correction

#### \* Within-subject: Coregistration

- Coregistration by maximising mutual information
- \* Between-subject: Normalise/Segment
- \* Between-subject: Dartel
- \* Smoothing

## Coregistration

- Inter-modal registration.
- Match images from same subject but different modalities:
  - anatomical localisation of single subject activations
  - achieve more precise spatial normalisation of functional image using anatomical image.





#### **Coregistration maximises Mutual Information**





Final Joint Histogram

T1 weighted

- \* Used for between-modality registration
- Derived from joint histograms
- \* MI=  $\int_{ab} P(a,b) \log_2 [P(a,b)/(P(a) P(b))]$ 
  - \* Related to entropy: MI = -H(a,b) + H(a) + H(b)
    - \* Where H(a) =  $-\int_a P(a) \log_2 P(a)$  and H(a,b) =  $-\int_a P(a,b) \log_2 P(a,b)$

#### "Check Reg" to assess alignment



















#### EPI dropout and distortion



#### References

- \* Friston et al. Spatial registration and normalisation of images. Human Brain Mapping 3:165-189 (1995).
- \* Collignon et al. Automated multi-modality image registration based on information theory. IPMI'95 pp 263-274 (1995).
- \* Thévenaz et al. Interpolation revisited. IEEE Trans. Med. Imaging 19:739-758 (2000).
- \* Andersson et al. *Modeling geometric deformations in EPI time series*. Neuroimage 13:903-919 (2001).
- \* Hutton et al. Image distortion correction in fMRI: a quantitative evaluation. NeuroImage 16:217-240 (2002).

#### **Pre-processing overview**

Statistics or whatever



#### **Alternative pipeline**





#### Contents

- \* Preliminaries
- \* Within-subject: Realignment
- \* Within-subject: EPI Distortion Correction
- \* Coregistration

#### \* Between-subject: Normalise/Segment

Use segmentation routine for spatial normalisation

- \* Gaussian mixture model
- \* Intensity non-uniformity correction
- \* Deformed tissue probability maps
- \* Between-subject: Dartel
- \* Between-subject: Smoothing

#### **Spatial normalisation**

- \* Brains of different subjects vary in shape and size.
- \* Need to bring them all into a common anatomical space.
  - \* Examine homologous regions across subjects
    - \* Improve anatomical specificity
    - \* Improve sensitivity
  - \* Report findings in a common anatomical space (eg MNI space)
- \* In SPM12, alignment is achieved by matching grey matter with grey matter and white matter with white matter.
  - \* With possible alignment of other tissues.

# Normalise/Segment

- \* This is the same algorithm as for tissue segmentation.
- \* Combines:
  - \* Mixture of Gaussians (MOG)
  - \* Bias Correction Component
  - \* Warping (Non-linear Registration) Component



# **Spatial normalisation**

- Default spatial normalisation in SPM12 estimates nonlinear warps that match tissue probability maps to the individual image.
- Spatial normalisation achieved using the inverse of this transform.



#### Segmentation

- Segmentation in SPM12 also estimates a spatial transformation that can be used for spatially normalising images.
- \* It uses a generative model, which involves:
  - Mixture of Gaussians (MOG)
  - \* Warping (Non-linear Registration) Component
  - \* Bias Correction Component





#### Modelling tissue intensities

 Classification is based on a *Mixture of Gaussians* model (MOG), which represents the intensity probability density by a number of Gaussian distributions.



Image Intensity —

## Tissue probability maps in SPM12

Includes additional non-brain tissue classes (bone, and soft tissue)

























































#### **Modelling deformations**



#### Modelling a bias field





**Corrupted image** 

**Bias Field** 

**Corrected image** 

#### Iterative optimisation scheme



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- \* Preliminaries
- \* Within-subject: Realignment
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- \* Within-subject: Coregistration
- \* Between-subject: Normalise/Segment

#### \* Between-subject: Dartel

- \* Velocity field parameterisation
- \* Objective function
- \* Template creation
- \* Examples
- \* Between-subject: Smoothing

# Dartel image registration

- \* Uses fast approximations
  - Deformation integrated using scaling and squaring
- \* Uses Levenberg-Marquardt optimiser
  - \* Multi-grid matrix solver
- Matches GM with GM, WM with WM etc
- Diffeomorphic registration takes about 30 mins per image pair (121×145×121 images).





Grey matter template warped to individual



Individual scan

#### Scaling and squaring example



# **Registration objective function**

\* Simultaneously minimize the sum of:

#### \* Matching Term

- Drives the alignment of the images.
- \* Multinomial assumption

#### \* Regularisation term

- \* A measure of deformation roughness
- \* Keeps the warps spatially smooth.
- \* A balance between the two terms.

#### Effect of different forms of regularisation





# Simultaneous registration of GM to GM and WM to WM



#### Template

Initial



















Average



Iteratively generated from 471 subjects

Began with rigidly aligned tissue probability maps

After a few iterations







**Final** template









Grey matter average of 452 subjects – affine







#### Grey matter average of 471 subjects

















































Subject 1









Subject 3





Evaluations of nonlinear registration algorithms



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- \* Between-subject: Normalise/Segment
- \* Between-subject: Dartel
- \* Between subject: Smoothing
  - Compensating for inaccuracies in inter-subject alignment

#### Smooth

#### Blurring is done by convolution.

Each voxel after smoothing effectively becomes the result of applying a weighted region of interest (ROI).



Before convolution



#### Convolved with a circle



#### Convolved with a Gaussian



#### Smooth



#### References

- \* Ashburner & Friston. Unified Segmentation. NeuroImage 26:839-851 (2005).
- \* Ashburner. A Fast Diffeomorphic Image Registration Algorithm. NeuroImage 38:95-113 (2007).
- \* Ashburner & Friston. Computing average shaped tissue probability templates. NeuroImage 45(2): 333-341 (2009).
- \* Klein et al. Evaluation of 14 nonlinear deformation algorithms applied to human brain MRI registration. NeuroImage 46(3):786-802 (2009).