# Function-Structure Integration in FreeSurfer



# Outline

- Function-Structure Integration
- Function-Structure Registration in FreeSurfer
- fMRI Analysis
  - Preprocessing
  - First-Level Analysis
  - Higher-Level (Group) Analysis
  - Correction for Multiple Comparisons
  - Data Hierarchies
- FreeSurfer Functional Analysis STream (FSFAST)
- Tutorial
- Demos

# **Function-Structure Integration**

- Viewing Functional Maps on Structural – Volume, Surface
- Inter-Subject Registration
- Region of Interest (ROI) Analysis
- Retinotopy
- Structural-Functional Covariates
  - Eg, use thickness at a voxels as covariate
  - Voxel-wise design matrices

# **FreeSurfer Registration**

#### FreeSurfer Subject-Specific

- Volumes
- Surfaces
- Thickness
- ROIs



#### Your Data/Software

- fMRI (FSL, SPM,...)
- DTI
- PET
- EEG/MEG ...

#### **Registration Matrix**

- Affine 4x4
- As many as 12 DOF (usually 6)
- Text file

# Registration

#### FreeSurfer Anatomical



Note: Registering the template functional volume to the anatomical volume is sufficient to register the template to the surface.

#### **Template Functional**

# Manual Registration

- tkregister2
- •Visually inspect registration
- Manually edit registration (6 DOF)
- Cf Manual Talairach registration





#### tkregister2 --help

# Tips

- Rigid = 6 DOF = No stretching
- Use CSF to get a sense of where the folds are
- Avoid using B0 distortion regions
- Avoid using ventricles
- Warning about "edge" of the brain
- Same Subject, Left-Right Flips





# **Command-line Tools**

#### Automatic Registration:

- fslregister help
- spmregister –help
  - FreeSurfer Scripts
- reg-feat2anat -help -

# Manual Registration:

• tkregister2 --help

#### **Transformations:**

- mri\_vol2surf --help
- mri\_vol2vol --help
- mri\_label2vol --help
- mri\_surf2vol --help

## Sampling on the Surface







- White/Gray
- Pial
- Half Way
- Average

Projection Fraction --projfrac 0.5

# Sampling on the Surface









fMRI Analysis Pipeline Overview



#### **fMRI** Preprocessing Stages

- Motion Correction
- Slice-timing Correction (Interleaved vs Seq)
- B0 Distortion Correction
- Intensity Normalization: 4D or 3D?
- Masking zeroing non-brain
- Resampling to Common Space
- Spatial Smoothing 3D or 2D?
- <u>Temporal Filtering is NOT Preprocessing!</u>

#### **Reasons for Smoothing**

- Improve CNR/SNR
- Reduce interpolation effects
- Make statistics more valid (GRF)
- Improve inter-subject registration
- Improve function-surface registration

## **Effects of Smoothing**

#### No Smoothing FWHM = 5mm



## **Effects of Smoothing**





#### **First Level Design and Analysis**

- First-Level = First Standard Deviation
- First-Level Design
  - Event Definition and HRF Specification
  - Nuisance Regressors
  - Temporal Filtering
  - Temporal Whitening
- First-Level Contrasts
  - Univariate (t) Pass up to next level
  - Multivariate (F)
- Analysis (Voxel-wise = "Massively Univariate")
  - Contrasts of HRF Amplitudes
  - Variances of the Contrasts

#### First Level Design: HRF Shapes



## Stimulus Schedule/FSFAST Paradigm File

- Codes Stimulus Schedule (and Weight)
- Four Columns
  - 1. Onset Time (Since Acq of 1<sup>st</sup> Saved Volume)
  - 2. Stimulus Code (0, 1, 2, 3...)
  - 3. Stumulus Duration
  - 4. Stimulus Weight (default is 1)
  - 5. Any other columns ignored
- Simple Text File
- Code 0 Always Fixation/NULL

0.000	0	15	1	Fixation
15.000	1	15	1	Task-Odd
30.000	0	15	1	Fixation
45.000	2	15	1	Task-Even
60.000	0	15	1	Fixation
75.000	1	15	1	Task-Odd
90.000	0	15	1	Fixation
105.000	2	15	1	Task-Even
120.000	0	15	1	Fixation
135.000	1	15	1	Task-Odd
150.000	0	15	1	Fixation
165.000	2	15	1	Task-Even
180.000	0	15	1	Fixation
195.000	1	15	1	Task-Odd
210.000	0	15	1	Fixation
225.000	2	15	1	Task-Even
240.000	0	15	1	Fixation

#### **First-Level Design Matrix**





**FIR** 

- Task convolved with HRF
- Polynomial (0-2) Nuisance Regressors
- MC Parameters reduced from 6 to 3



#### Higher-Level (Group) Analysis

- Higher-Level Design
  - Groups and covriates
  - Contrasts
- Analysis Method
  - Random Effects (RFx, OLS = ordinary least squares)
  - Weighted Random Effects (WRFx, WLS=weighted least squares)
  - Mixed Effects
  - Fixed Effects (FFx)
- Correction for Multiple Comparisons
  - Clustering (GRF, Monte Carlo, Permutation)

#### **Group Effect Models**

- Random Effects (RFx, OLS; WRFx, WLS)
  - Does effect exist in the general population that my subjects were drawn from?
  - Weighted weight each subject by 1/First Level Noise
- <u>Fixed Effects</u> (FFx) Does effect exist within the group of subjects that I am studying? Like having one subject scanned multiple times.
- <u>Mixed Effects</u> use First Level (within-subject) Noise AND between-subject noise to do better weighting.

#### **One-Sample Group Mean (OSGM)**

- No groups, No Covariates
- Does average = 0?
- One-sample t-test
- Group Design Matrix: Vector of All 1s

#### **FS-FAST** Directory Hierarchy



Use unpacksdcmdir to import Session in Siemens dicom to FS-FAST.

#### **FS-FAST** Tutorial

#### • Data - fBIRN

- 5 Subjects
- 4 Runs Each (TR=3, 85TP)
- Sensory Motor Task
- 15 sec Blocks
- 9 OFF
- 8 ON
- Code Odd and Even Separately
- Test Odd vs Even

# fBIRN Phase I Sensory-Motor Task Schedule

#### surfer.nmr.mgh.harvard.edu/fswiki/FsFastTutorial

## **FS-FAST** Tutorial Exercises

- Data setup
  - "Import" in to hierarchy
  - Create paradigm files
  - Link to FreeSurfer Anatomical Analysis
- Viewing Functional Results in TkMedit/TkSurfer
- Preprocessing MC and Smoothing
- Registration automated and manual
- First Level
  - Design and Contrasts: Gamma, Finite Impulse Response (FIR)
  - First Level Analysis
  - Visualization volume and surface
- Group Level Analysis One-Sample Group Mean (OSGM)
  - QA
  - RFx, WRFx, FFx
  - Volume (Talairach) and Surface

#### **FS-FAST** Tutorial Exercises

- Four main directories at various levels of processing in \$FSFTUTDIR:
  - 1. <u>fb1-raw</u> raw data, nifti format, unorganized
  - 2. <u>fb1-raw-study</u> raw data organized in FSFAST hierarchy
  - 3. <u>fb1-preproc-study</u> preprocessed data
  - 4. <u>fb1-analysis-study</u> fully analyzed
    - 1. First-level Analyses
    - 2. Group Analyses in Tal and Surf
- You don't necessarily need to run any processing can just run visualization.

Start Terminal firefox& surfer.nmr.mgh.harvard.edu/fswiki/FsFastTutorial