

# Practice in resting-state fMRI (rs-fMRI) Analysis: PART I

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## Course Arrangement

### PART I (10/3)

- rs-fMRI pre-processing
- REST and DPARSF
- REST go through: ReHo, Functional Connectivity, ALFF, fALFF, utilities

### PART II (10/4)

- Advanced connectivity analysis
- DPARSF go through
- Statistics



## Download Demo Materials

心智科學腦研究推動網

心智影像研究(MRI)中心 @成大 活動網頁 → 實作資料

[http://fmri.ncku.edu.tw/tw/course\\_view.php?no=126](http://fmri.ncku.edu.tw/tw/course_view.php?no=126)

盧家鋒 個人網頁

靜息態功能性磁振影像分析實作 → 實作資料

[http://www.ym.edu.tw/~cflu/CFLu\\_course\\_speech.html](http://www.ym.edu.tw/~cflu/CFLu_course_speech.html)

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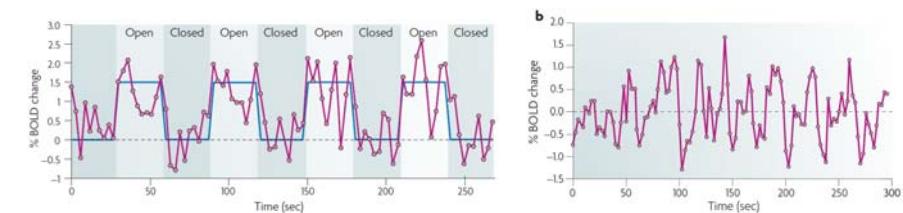
## Spontaneous Fluctuation

### Task-specific fMRI

- ✓ Model-based Analysis
- ✓ Model-free (data-driven) Analysis

### Resting-state fMRI (rs-fMRI)

- ✓ Model-free (data-driven) Analysis



*Nature Reviews Neuroscience* 8.9 (2007): 700-711.

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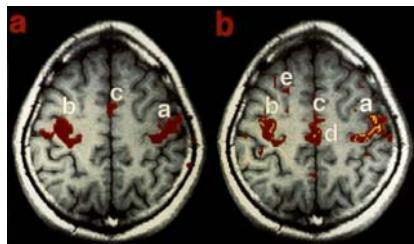
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# First rs-fMRI Article (MRM 1995)



## Functional Connectivity in the Motor Cortex of Resting Human Brain Using Echo-Planar MRI

Bharat Biswal, F. Zerrin Yetkin, Victor M. Haughton, James S. Hyde



a. Functional activation during tasking  
b. rs-fMRI correlation maps  
(red: positive, yellow: negative)

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## Rs-fMRI Analyses

### 1. Functional connectivity analysis

- Linear correlation
- Granger causality analysis (GCA), effective connectivity
- Independent component analysis (GIFT-ICA; Calhoun et al., NeuroImage 2001)

### 2. Depicting local features of BOLD signal

- Regional homogeneity (ReHo; Zang et al., NeuroImage 2004)
- Amplitude of low-frequency fluctuation (ALFF; Zang et al., Brain & Development 2007)
- Fractional ALFF (fALFF; Zou et al., J Neurosci Methods 2008)

Functional Integration ⇔ Functional segregation

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## Preprocessing for rs-fMRI

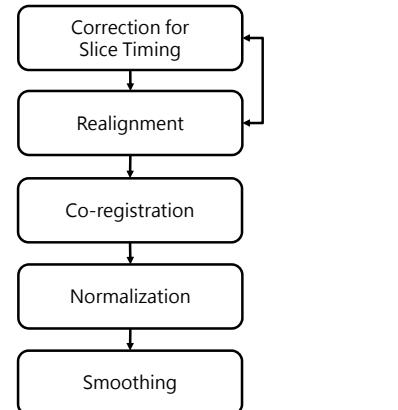
Similar to that used for task-specific fMRI analysis

- Slice timing
- Realignment
- Co-registration (with anatomical images)
- Normalization
- Smoothing
- Segment (tissue classification; optional)



Standard procedure for fMRI preprocessing

[http://www.ym.edu.tw/~cflu/CFLU\\_course\\_mriprep.html](http://www.ym.edu.tw/~cflu/CFLU_course_mriprep.html), Week 16



## Switch current folder to data folder

More convenient to execute the subsequent processing steps....

Current Folder: C:\Users\Alvin\Desktop\data\data\Subj01

data	
名稱	類型
Subj01	檔案資料夾
Subj02	檔案資料夾
Subj03	檔案資料夾

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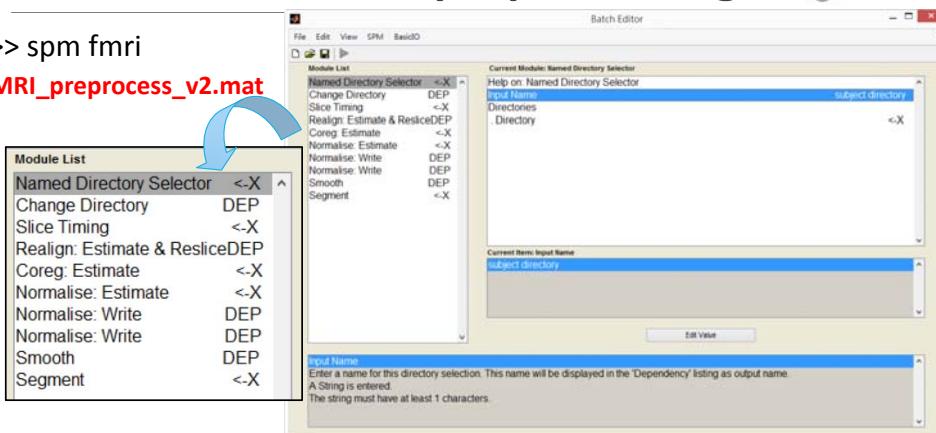
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# Batch of SPM fMRI preprocessing



>> spm fmri

**fMRI\_preprocess\_v2.mat**



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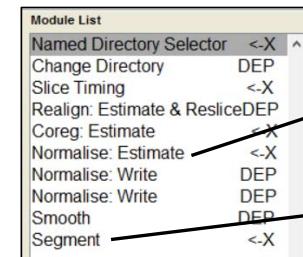
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# Batch of SPM fMRI preprocessing



>> spm fmri

**Save as your own batch template**



**T1 template image**

C:\Users\Alvin\Desktop\softwares\spm8\templates\T1.nii,1

**Tissue probability maps**

C:\Users\Alvin\Desktop\softwares\spm8\tpm\grey.nii,1

C:\Users\Alvin\Desktop\softwares\spm8\tpm\white.nii,1

C:\Users\Alvin\Desktop\softwares\spm8\tpm\csf.nii,1

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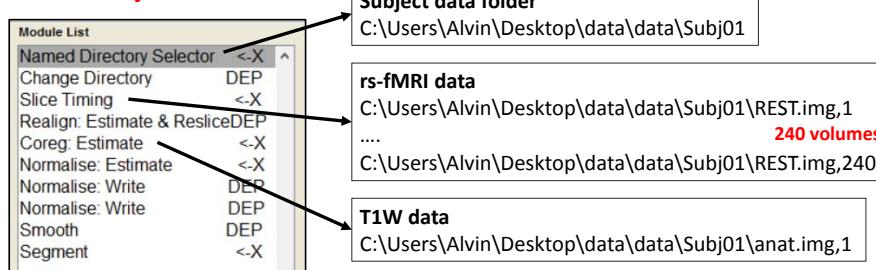
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# Batch of SPM fMRI preprocessing



>> spm fmri

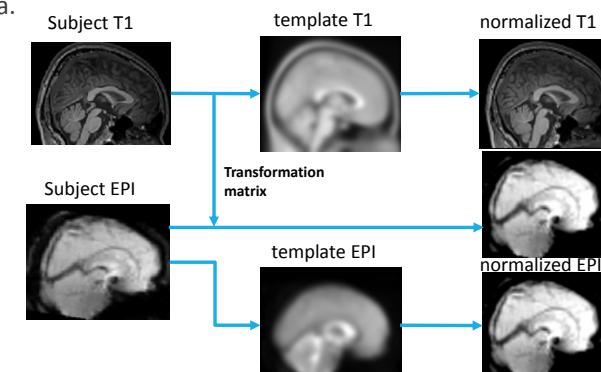
**Save as subject's batch**



# Normalization



We can perform spatial normalization using either anatomical (T1) images or fMRI (EPI) data.



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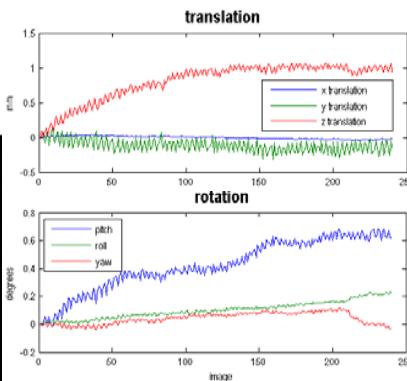
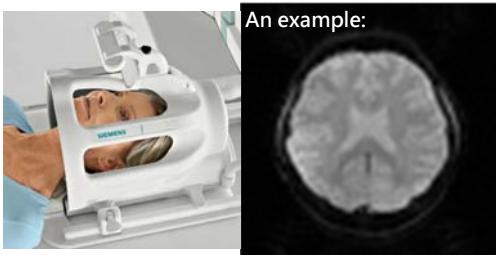
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## Realignment of head motion



The signal variation from movement is larger than hemodynamic response.

6-parameter Rigid body registration & transformation (align to the 1<sup>st</sup> volume)  
→ 6 co-variates for rs-fMRI analysis



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## Employed Software/Package



### 1. SPM preprocessing

- <http://www.fil.ion.ucl.ac.uk/spm/>

### 2. REST functional connectivity, ReHo, ALFF, fALFF, VMHC

- <http://restfmri.net/forum/index.php?q=rest>

### 3. DPARSF/DPABI

- <http://fmri.org/DPARSF>

- Data Processing Assistant for Resting-State fMRI (DPARSF)

- Based on SPM and REST toolbox

REST: Song et al., PLoS ONE, 2011.

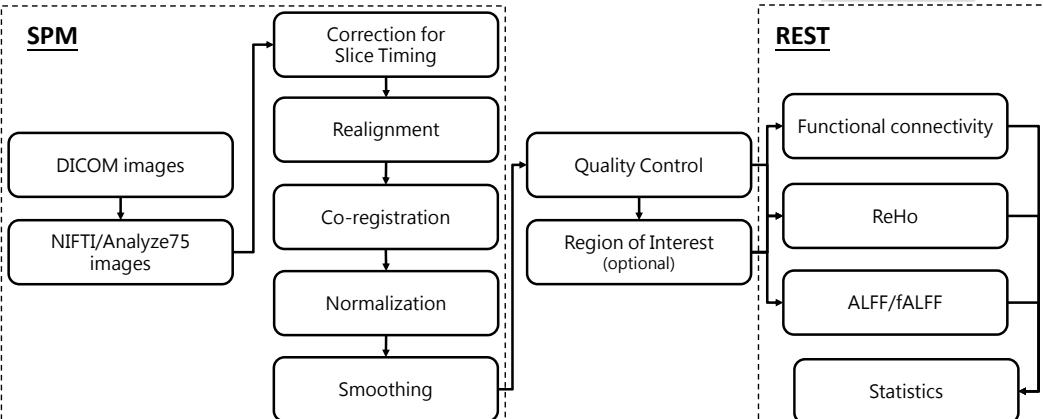
DPARSF: Yan et al., Frontiers in System Neuroscience, 2010.

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## Processing Flow in DPARSF



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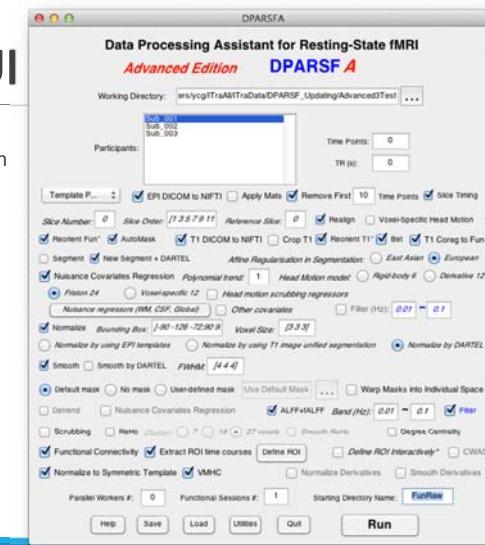
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## DPARSF GUI



### One-stop service

- Similar to the SPM batch
- Easy to process, but...
- Obscure to novice : (



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## Easier Way to Learn : )



SPM8 (Alvin): Menu

SPM+REST

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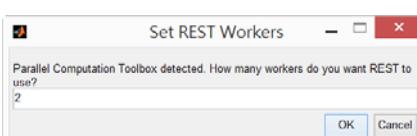
HTTP://WWW.YM.EDU.TW/~CFLU

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## Initialize REST environment



>> rest



Welcome: Alvin, 20151002\_2312  
REST Version: 1.8, Release: 20130615  
Citation Information:  
Xiao-Wei Song, Zhang-Ye Dong, Xiang-Yu Long, Su-Fang Li, Xi-Nian Zu  
Starting matlabpool using the 'local' profile ... connected to 2 workers.  
Now REST is Running on 2 workers.

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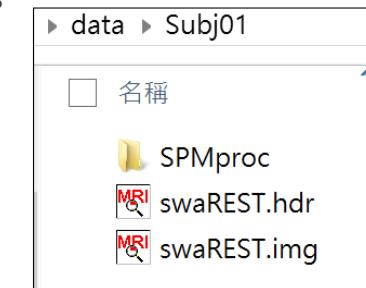
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## Organize data folder



Before using REST toolbox,

Create a new subfolder to archive all SPM-processed data except the file pair of "swaREST.hdr/img"



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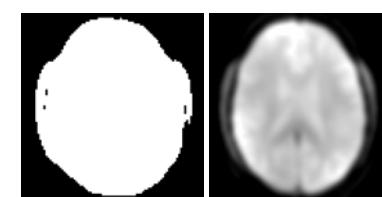
HTTP://WWW.YM.EDU.TW/~CFLU

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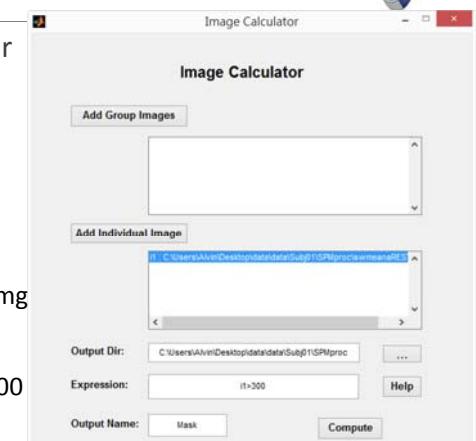
## Create Brain Mask



Utilities → REST Image Calculator



i1 > 300



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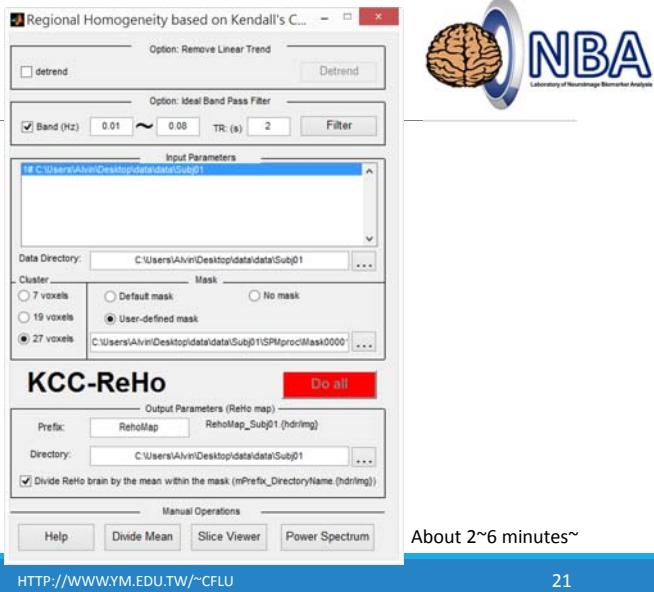
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## ReHo Setup

Assign folder with Analyze75 image pair (.img/.hdr), smoothed images

Select the user-defined mask

Select output directory and prefix



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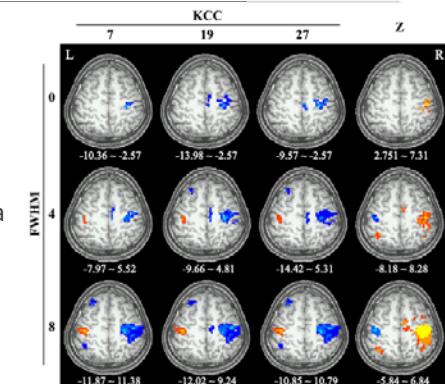
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## What does ReHo represent?

Regional homogeneity, temporal similarity between neighbor voxels within a small cluster.

"ReHo supposed that voxels within a functional brain area were more temporally homogeneous when this area is involved in a specific condition."

Model-free, data-driven ReHo can reflect cortical activation.



Zang et al., NeuroImage 2004.

## An example of ReHo (KCC)



		OBJECTS					n=5	
		i: 5 time points (number of ranks)						
JUDGES		rank	4	3	1	5	2	
j: 3 voxels within a cluster	K=3		4	3	1	5	2	
			4	3	1	5	2	

$$R_1=12, R_2=9, R_3=3, R_4=15, R_5=6,$$

$$\bar{R} = 9, S = 90$$

$$W = \frac{12 \times 90}{9(125 - 5)} = 1 \quad \rightarrow \text{unanimous measurements}$$

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## An example of ReHo (KCC)



		OBJECTS					n=5	
		i: 5 time points (number of ranks)						
JUDGES		rank	4	3	1	5	2	
j: 3 voxels within a cluster	K=3		5	1	2	3	4	
			3	2	4	1	5	

$$R_1=12, R_2=6, R_3=7, R_4=9, R_5=11,$$

$$\bar{R} = 9, S = 24$$

$$W = \frac{12 \times 24}{9(125 - 5)} = 0.27 \quad \rightarrow \text{Lower concordance}$$

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## An example of ReHo (KCC)



		OBJECTS				
		i: 5 time points (number of ranks)				
		n=5				
JUDGES		4	3	1	5	2
j: 3 voxels within a cluster		3	5	4	1	2
K=3		2	1	4	3	5

$$R_1=9, R_2=9, R_3=9, R_4=9, R_5=9,$$

$$\bar{R} = 9, S = 0$$

$$W = \frac{12 \times 0}{9(125 - 5)} = 0 \quad \rightarrow \text{Total disagreement}$$

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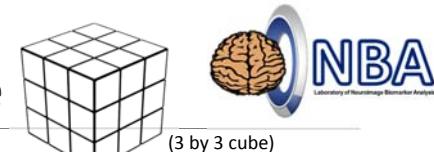
$$R_i = \sum_{j=1}^K r_{i,j}$$

$$\bar{R} = \frac{1}{n} \sum_{i=1}^n R_i$$

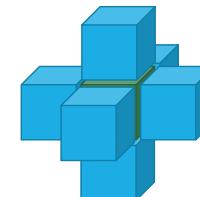
$$S = \sum_{i=1}^n (R_i - \bar{R})^2$$

$$W = \frac{12S}{K^2(n^3 - n)}$$

## Neighbors & Cluster size

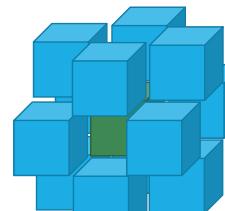


Surface connected (6)



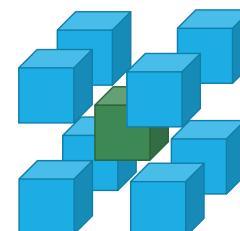
6+1= 7-voxel cluster

Edge connected (12)



6+12+1= 19-voxel cluster

Corner connected (8)



6+12+8+1= 27-voxel cluster

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## Computation time: 2 ~6 mins



```
Ideal rectangular filter: "C:\Users\Alvin\Desktop\NCKU-REST\data\Subj01"
Read 3D EPI functional images: "C:\Users\Alvin\Desktop\NCKU-REST\data\Subj01".
Load mask "".
Band Pass Filter working. Wait.....
Saving filtered images. Wait...
Band pass filter over.
Elapsed time is 194.178927 seconds.
```

ReHo:

```
Read these 3D EPI functional images. wait...
Read 3D EPI functional images: "C:\Users\Alvin\Desktop\NCKU-REST\data\Subj01_filtered".
Rank calculating.....
Calculate the kcc on voxel by voxel for the data set.....
Regional Homogeneity computation over, elapsed time: 167.781 seconds
```

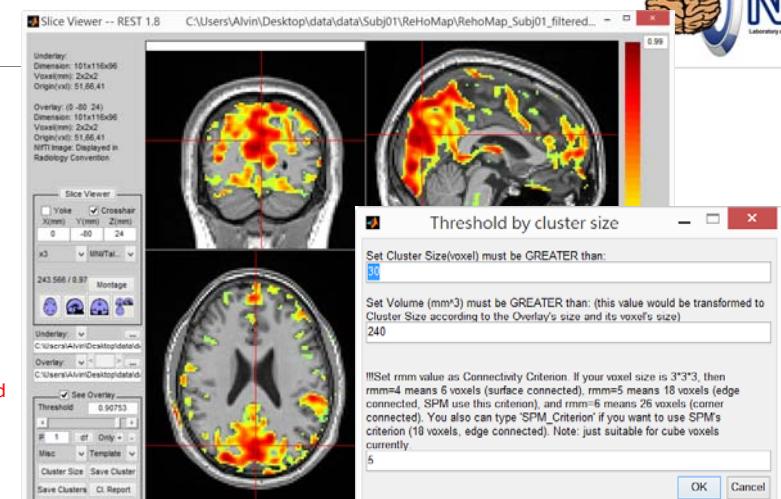
## ReHo Slice Viewer



Underlay:  
manat.img

overlay:  
RehoMap\*.nii

Adjust threshold



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# Functional Con.

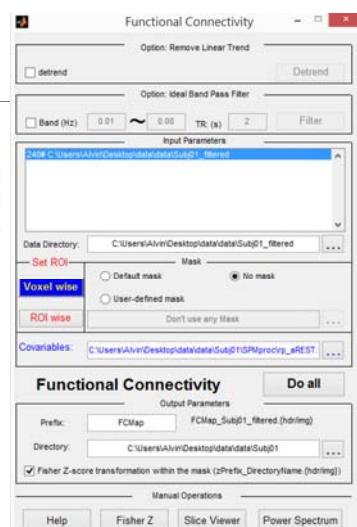
Select folder of Subj01\_filtered  
(No need to apply filter and  
mask again)

Voxel wise ⇔ ROI wise

6 motion parameters as  
covariates

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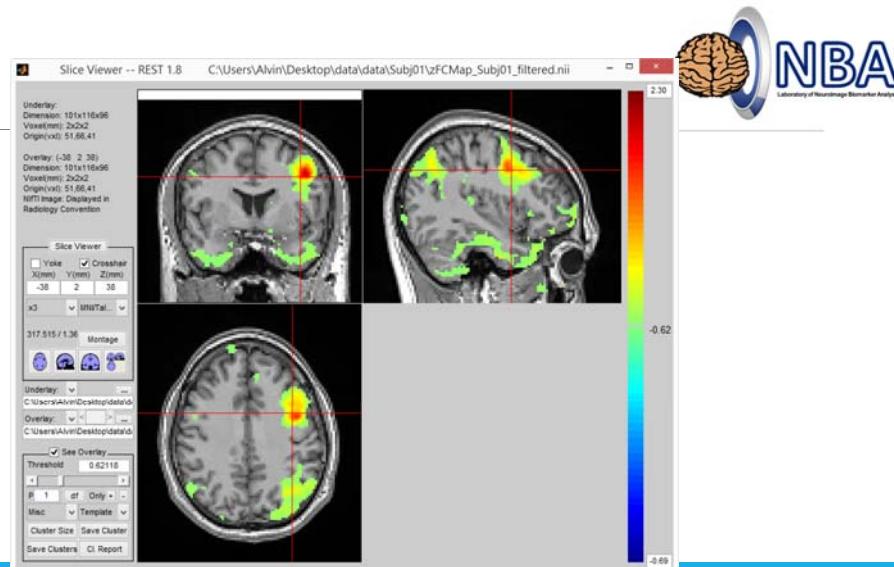
# Sphere ROI



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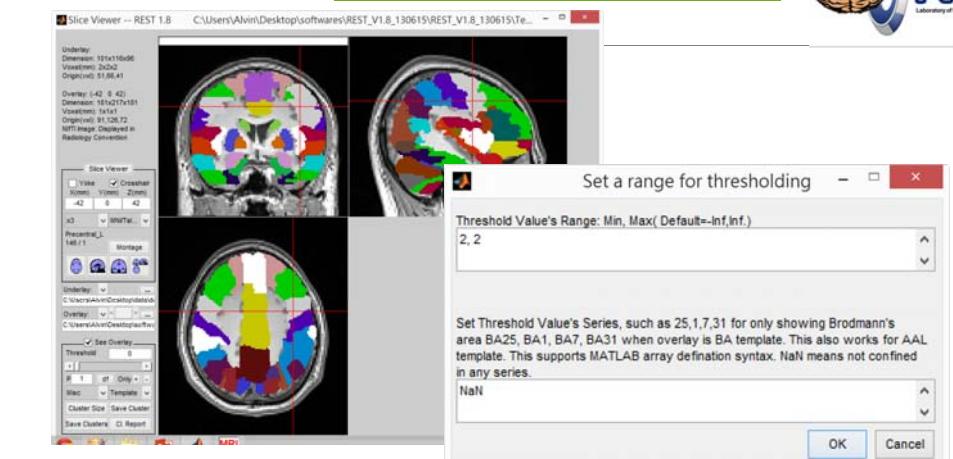
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# Atlas ROI

IBASPM 64-bit version  
[http://www.ym.edu.tw/~cflu/software/lbaspm\\_64.zip](http://www.ym.edu.tw/~cflu/software/lbaspm_64.zip)



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## What does ALFF represent?

Regional activation (regional activity during resting state).

Regional spontaneous neural activity

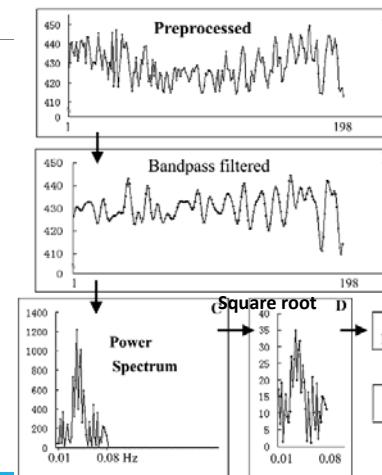
- ALFF is higher in grey matter than in white matter (Biswal et al., 1995).
- Kiviniemi et al. [22] reported activation in the visual cortex due to low-frequency fluctuations at about 0.034 Hz using the power spectrum method.

ReHo and functional connectivity analyses focus on the similarities of intra- and inter-regional time series, respectively, and ALFF measures the amplitude of regional activity.

Zang et al., Brain & Development 2007



## ALFF ALFF flowchart



BOLD time series for a given voxel

Zang et al., Brain & Development 2007



## Fractional ALFF

However, it has been indicated that the ALFF is also sensitive to the physiological noise.

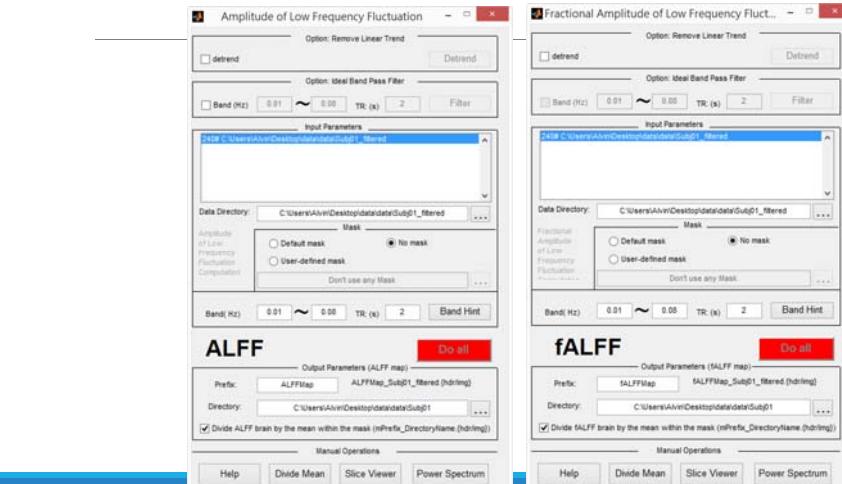
A fractional ALFF (fALFF) approach, i.e., the ratio of power spectrum of low-frequency (0.01–0.08 Hz) to that of the entire frequency range.

The non-specific signal components in the cistern areas in resting-state fMRI were significantly suppressed, indicating that the fALFF approach improved the sensitivity and specificity in detecting spontaneous brain activities.

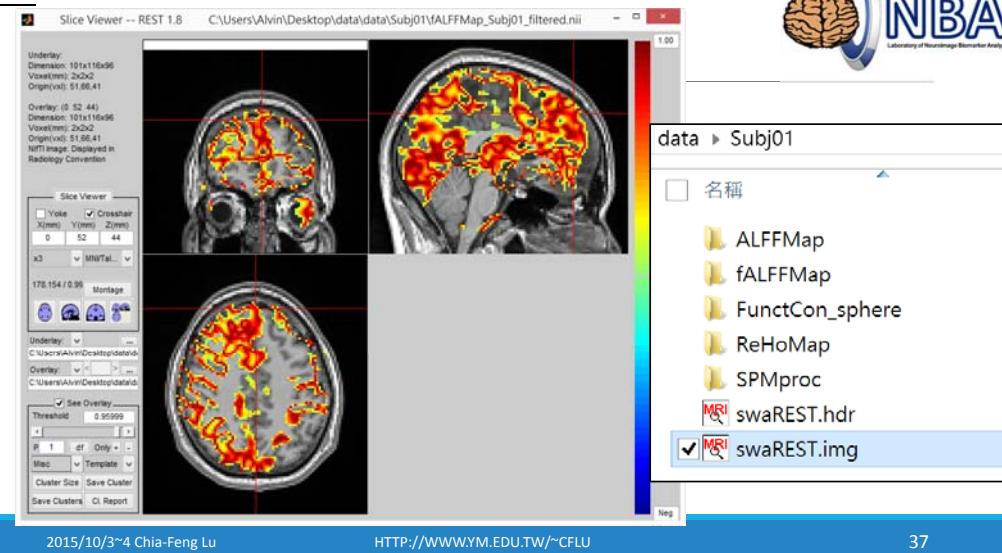
Zou et al., J Neurosci Methods 2008



## ALFF & ALFF ALFF/fALFF Setup



fALFF

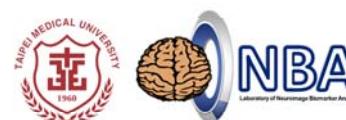


## Q & A

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Translational Imaging Research Center  
Taipei Medical University

TIRC Team  
2015.7.20 at TMU

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