

# Advanced Analyses of Resting-State fMRI (rs-fMRI): PART I

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## 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/download/20160528\\_29rsfMRI/adrsfMRIdata.zip](http://fmri.ncku.edu.tw/tw/download/20160528_29rsfMRI/adrsfMRIdata.zip)

盧家鋒 個人網頁

進階fMRI資料分析理論與實作 → 實作資料

<http://www.ym.edu.tw/~cflu/adrsfMRIdata.zip>

## Course Arrangement

### PART I (5/28)

- Preprocessing of rs-fMRI
- SPM preprocessing batch
- Modification of batch file
- Group independent component analysis (group ICA)

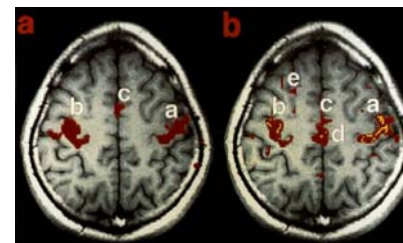
### PART II (5/29)

- Seed-based functional connectivity
- Complex network analysis

## 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)

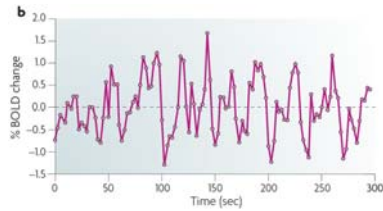
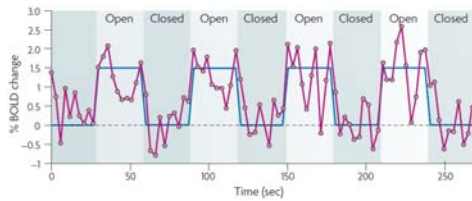
# Spontaneous Fluctuation

## 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.

# Rs-fMRI Analyses

## 1. Functional connectivity analysis

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

## 2. Depicting local features of BOLD signal

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

- 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

# 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>

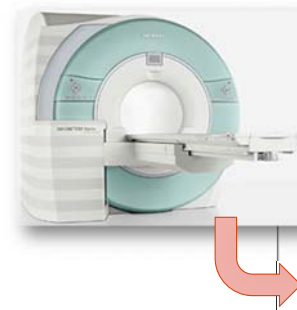
## 3. GIFT (v3.0a), group ICA

- <http://mialab.mrn.org/software/gift/#>

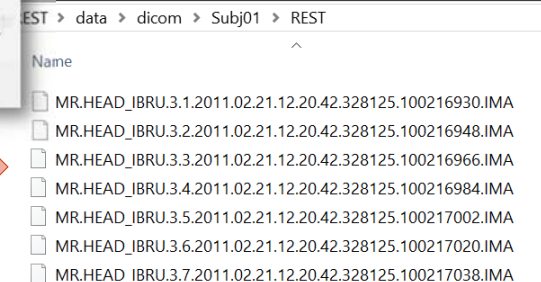
## 4. Brain Connectivity Toolbox/Network Based Statistic Toolbox

- <https://sites.google.com/site/bctnet/>

# Let's Start from Very Beginning



**DICOM format!!** ↔ NIfTI (.nii) or Analyze75 (\*.hdr, \*.img)



# DICOM (1993)

- Digital Imaging and COmmunication in Medicine



NEMA, Suite 1752  
1300 North 17<sup>th</sup> Street  
Rosslyn, VA 22209  
Ph: (703) 841-3285  
<http://dicom.nema.org>

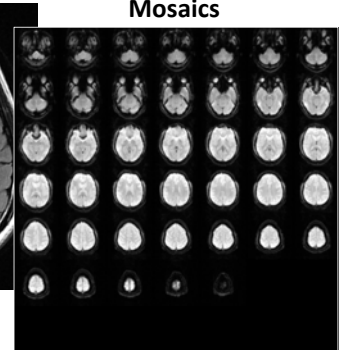
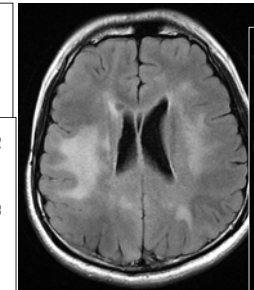
- ACR & NEMA formed a committee in 1983
  - American College of Radiology
  - National Electrical Manufacturers Association

**“The Common Language of Medical Equipment”**

# DICOM format

Header + Image data

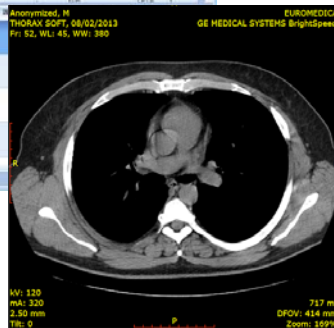
```
0008.0020.Study Date=20151204
0008.0021.Series Date=20151204
0008.0022.Acquisition Date=20151204
0008.0023.Image Date=20151204
0008.0030.Study Time=122630
0008.0031.Series Time=122800
0008.0032.Acquisition Time=122800
0008.0033.Imag0018,0050.Slice Thickness=5
0008.0050.Acc0018,0080.Repetition Time [TR, ms]=8002
0008.0060.Mod0018,0081.Echo Time [TE, ms]=127.948
0008.0070.Mar0018,0082.Inversion Time=2000
0008.0080.Inst0018,0083.Number of Averages=1
0008.0090.Ref0018,0084.Imaging Frequency=63.854903
0008.1010.Sta0018,0085.Imaged Nucleus=1H
0008.103E.Seri0018,0086.Echo Number=
0008.1070.Ope0018,0087.Magnetic Field Strength=1.5
0008.1090.Mar0018,0088.Spacing Between Slices=6
0018,0091.Echo Train Length=1
```



Retrieve of subject/imaging information.  
Use with caution! Personal privacy!

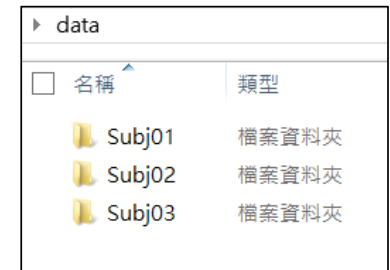
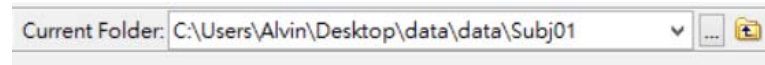


- Data query
- Image Information
- Communication

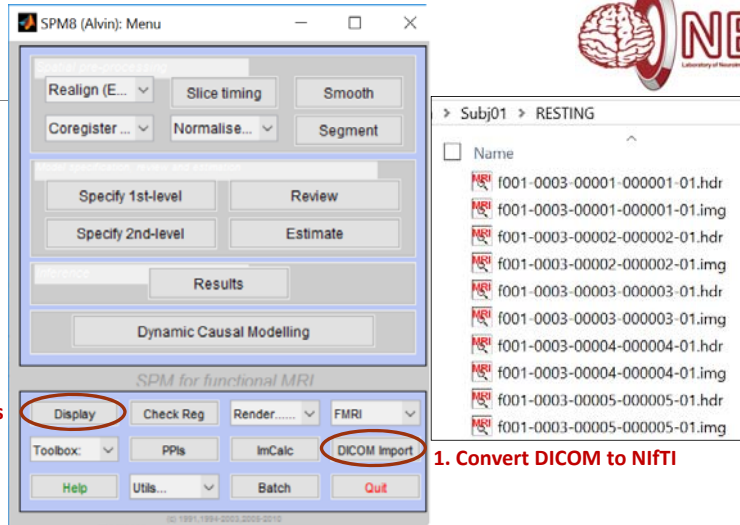


# Switch current folder to data folder

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



# DICOM import



2. Check converted images

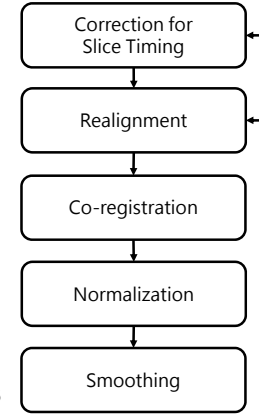
1. Convert DICOM to NIFTI



# 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)



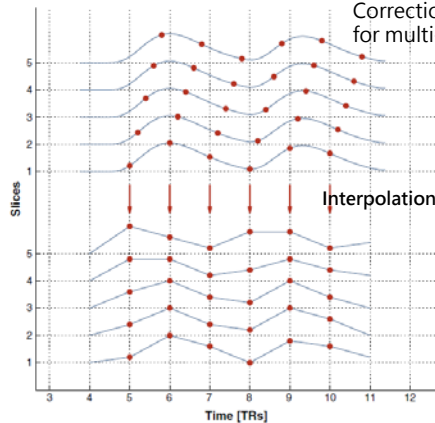
Standard procedure for fMRI preprocessing

[http://www.ym.edu.tw/~cflu/CFLu\\_course\\_mrjprinp.html](http://www.ym.edu.tw/~cflu/CFLu_course_mrjprinp.html), Week 16

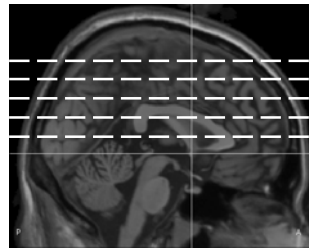
# Correction for Slice Timing



Correction for slightly different imaging timing for multi-slice acquisition in a TR.



For example:  
Acquire 5 slices in 1 TR  
→ Temporal offset between slices



Sladky et al, NeuroImage 2011,58:588-594.

# Realignment of head motion

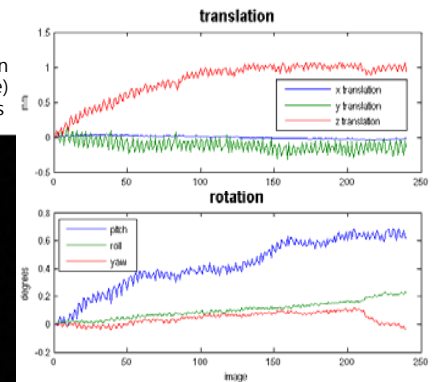
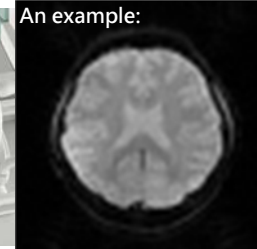


The signal variation from movement is larger than hemodynamic response.

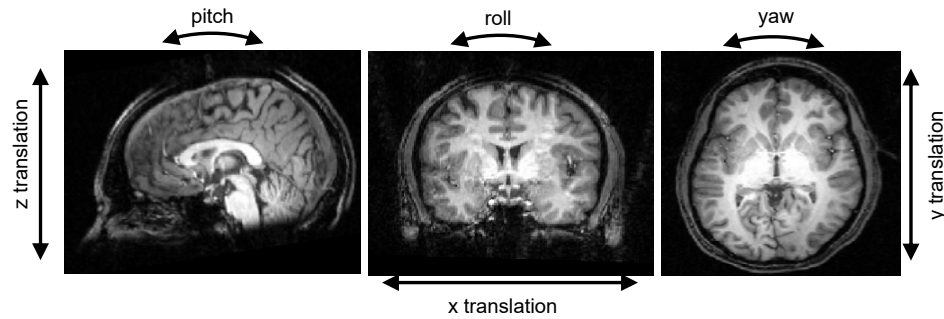
6-parameter Rigid body registration & transformation (align to the 1<sup>st</sup> volume)  
→ 6 co-variables for rs-fMRI analysis



An example:



## 3 translations and 3 rotations



fMRI slides from <http://culhamlab.ssc.uwo.ca/fmri4newbies/Tutorials.html>

## EPI undistortion/unwarp



Magnetic inhomogeneity can cause

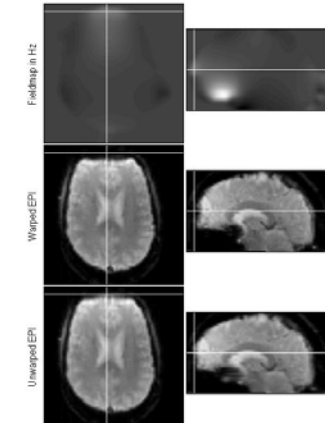
- Signal loss
- Spatial distortion

Magnetic field warps at tissue boundaries

- The frontal pole, orbito-frontal cortex
- Medial temporal lobe (hippocampus)

The benefit of undistortion/unwarp

- Make the shape of an individual's fMRI data more similar to their anatomical images.

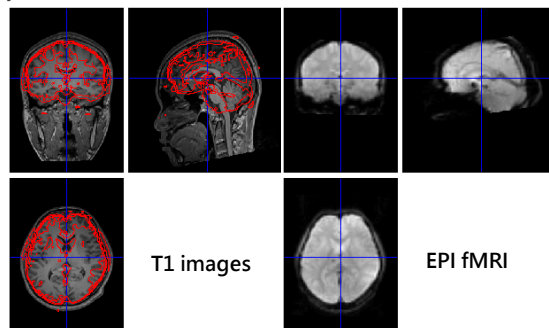


## Co-registration



Align fMRI (EPI) data to structural (T1) images.

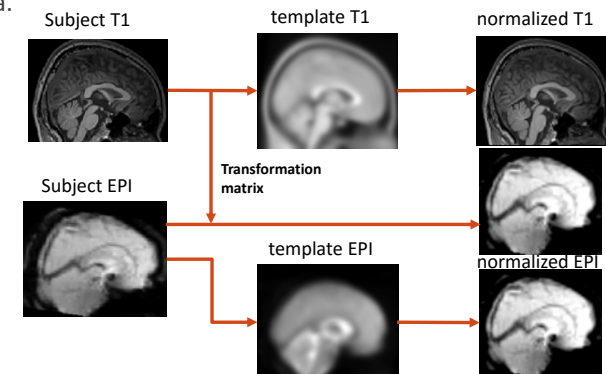
- Rigid body transformation using mutual information
- Manual adjustment



## Normalization

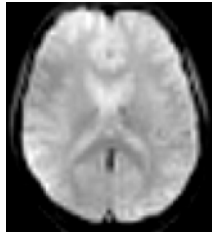


We can perform spatial normalization using either anatomical (T1) images or fMRI (EPI) data.

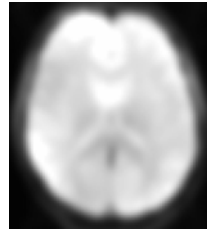


# Gaussian Smoothing

- Each voxel becomes weighted average of surrounding voxels.
- Render the data more normally distributed.
- Compensate for inaccuracies in normalization between individuals.
- Increase signal-to-noise ratio



normalization

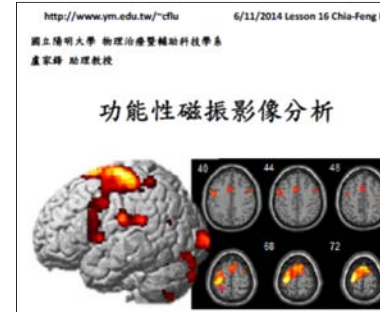


Normalization+smoothing

2 x 2 x 2 mm<sup>3</sup>

# Preprocessing Step-by-step

## fMRIprocess\_CFLu.pdf



http://www.ym.edu.tw/~cflu 6/11/2014 Lesson 16 Chia-Feng Lu

三、使用 SPM

1. 開啟 MATLAB (請確認已完成 Set Path), 在 command window 鍵入 `spm_fmri`
2. 依序練習執行 (請使用 dataset\_L16/Subj01 中的影像)
  - (1) Slice timing: 校正不同切片取樣時間的差異 (Siemens MRI 之 interleave images, 如為偶數張切片取樣順序為 2, 4, ..., 1, 3, ...; 奇數張切片順序為 1, 3, ..., 2, 4, ...)
  - (2) Realign (Est & Res): 校正不同 volume 間的頭部移動
  - (3) Coregister (Estimate): 將 T1 影像對位至 fMRI 影像 (使用 Check Reg 檢查對位結果)
  - (4) Segment: 計算 GM、WM 與 CSF 的機率圖形
  - (5) Normalise: 將影像標準化至 ICBM-152 模板 (Bounding box: -100 -130 -80; 100 100 110)
  - (4) Smooth: 對 fMRI 影像進行三維高斯平滑化處理

[http://www.ym.edu.tw/~cflu/MRI\\_PrinApp\\_Class16\\_CFLu.pdf](http://www.ym.edu.tw/~cflu/MRI_PrinApp_Class16_CFLu.pdf)

# Batch Step-by-step

## 五、附錄：Batch 檔設定詳細說明

- 建立基本處理流程(dataset\_L16fMRIbatch\_raw.mat)
1. 請先將下列順序的 module 加入至 batch 中
    - ✓ BasicIO -> Named Directory Selector
    - ✓ BasicIO -> Change Directory
    - ✓ BasicIO -> Make Directory
    - ✓ SPM -> Temporal -> Slice Timing
    - ✓ SPM -> Spatial -> Realign: Estimate & Reslice
    - ✓ SPM -> Spatial -> Coreg: Estimate
    - ✓ SPM -> Spatial -> Segmentation
    - ✓ SPM -> Spatial -> Normalise: Write
    - ✓ SPM -> Spatial -> Smooth

http://www.ym.edu.tw/~cflu 6/11/2014 Lesson 16 Chia-Feng Lu

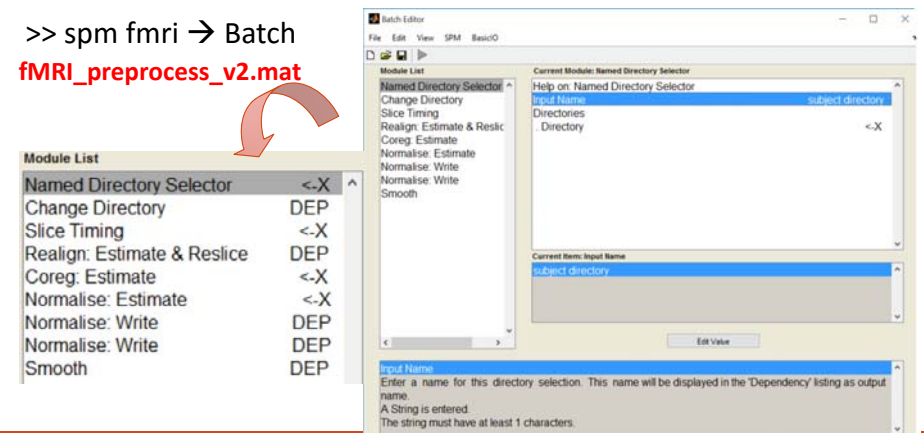
➢ 輸入基本參數(dataset\_L16fMRIbatch\_process.mat)

請按照以下步驟依序設定各 module 的參數

- ✓ **Named Directory Selector**: Input Name 填入 subject directory; 在 Directories 中點兩下讓 Directory 出現一個 <-X 的標記
- ✓ **Make Directory**: New Directory Name 輸入 stats, 提供 SPM 統計分析擺放檔案的位置
- ✓ **Slice Timing**: Data 點兩下出現 <-X; Number of Slices: 40; TR:2; TA 輸入 2-(2/40); Slice order:[2:2:40,1:2:40]; Reference Slice:1
- ✓ **Realign**: Data 點兩下, 讓 Session 出現 <-X
- ✓ **Normalise**: Data 點兩下加入一個 Subject; 更改 bounding box 為 -100 -130 -80; 100 100 110
- ✓ **Smooth**: 可調整 FWHM 參數

# Batch of SPM fMRI preprocessing

>> spm\_fmri -> Batch  
fMRI\_preprocess\_v2.mat



The screenshot shows the SPM Batch Editor window. On the left, a 'Module List' contains the following modules with checkboxes: Named Directory Selector (checked), Change Directory (DEP), Slice Timing (checked), Realign: Estimate & Reslice (DEP), Coreg: Estimate (checked), Normalise: Estimate (checked), Normalise: Write (DEP), Normalise: Write (DEP), and Smooth (DEP). A red arrow points from the text 'fMRI\_preprocess\_v2.mat' to the 'Named Directory Selector' module. On the right, the 'Current Module: Named Directory Selector' dialog is open, showing 'subject directory' selected in the 'Input Name' field and a '<-X' marker in the 'Directories' list.

# Batch of SPM fMRI preprocessing



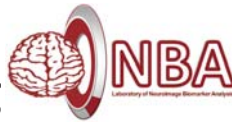
>> spm fmri

Save as your own batch template

| Module List                 |       |
|-----------------------------|-------|
| Named Directory Selector    | <-X ^ |
| Change Directory            | DEP   |
| Slice Timing                | <-X   |
| Realign: Estimate & Reslice | DEP   |
| Coreg: Estimate             | <-X   |
| Normalise: Estimate         | <-X   |
| Normalise: Write            | DEP   |
| Normalise: Write            | DEP   |
| Smooth                      | DEP   |

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

# Batch of SPM fMRI preprocessing



>> spm fmri

Save as subject's batch

| Module List                 |       |
|-----------------------------|-------|
| Named Directory Selector    | <-X ^ |
| Change Directory            | DEP   |
| Slice Timing                | <-X   |
| Realign: Estimate & Reslice | DEP   |
| Coreg: Estimate             | <-X   |
| Normalise: Estimate         | <-X   |
| Normalise: Write            | DEP   |
| Normalise: Write            | DEP   |
| Smooth                      | DEP   |

**Subject data folder**  
 C:\Users\Alvin\Desktop\data\data\Subj01

**rs-fMRI data**  
 C:\Users\Alvin\Desktop\data\data\Subj01\REST.img,1  
 .....  
 C:\Users\Alvin\Desktop\data\data\Subj01\REST.img,240  
**240 volumes**

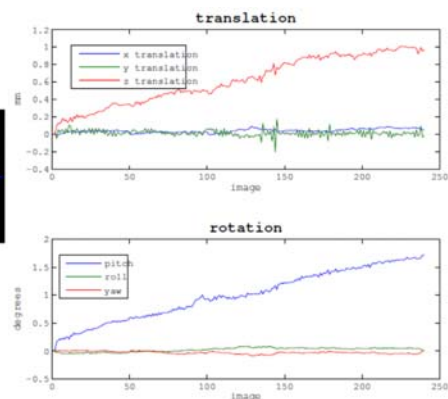
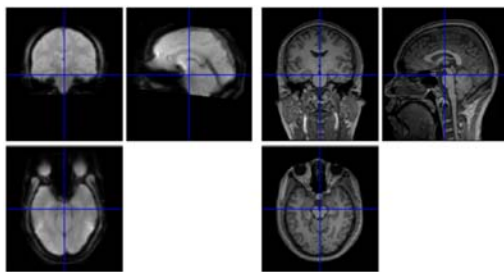
**T1W data**  
 C:\Users\Alvin\Desktop\data\data\Subj01\anat.img,1

# Always Check Your Data!



SPM processing logfile (spm\_2016May28.ps)

<https://online2pdf.com/convert-ps-to-pdf>



# Modification of Batch File



Batch Editor → View → Show .m Code

```
Batch Code Browser
matlabbatch(1).cfg_basico.cfg_named_dir.name = 'subject directory';
matlabbatch(1).cfg_basico.cfg_named_dir.dirs = {'C:\Users\Alvin\Desktop\data_proc\Subj02\'};
matlabbatch(2).cfg_basico.cfg_cd.dir(1) = cfg_dep;
matlabbatch(2).cfg_basico.cfg_cd.dir(1).tname = 'Directory';
matlabbatch(2).cfg_basico.cfg_cd.dir(1).tgt_spec(1)(1).name = 'filter';
matlabbatch(2).cfg_basico.cfg_cd.dir(1).tgt_spec(1)(1).value = 'dir';
matlabbatch(2).cfg_basico.cfg_cd.dir(1).tgt_spec(1)(2).name = 'strtype';
matlabbatch(2).cfg_basico.cfg_cd.dir(1).tgt_spec(1)(2).value = 'e';
matlabbatch(2).cfg_basico.cfg_cd.dir(1).sname = 'Named Directory Selector: subject directory(1)';
matlabbatch(2).cfg_basico.cfg_cd.dir(1).src_exbranch = substruct('.', 'val', '{', '{', 1, '-', 'val', '{', '{', 1});
matlabbatch(2).cfg_basico.cfg_cd.dir(1).src_output = substruct('.', 'dirs', '{', '{', 1});
%%
matlabbatch(3).spm_temporal.st.scans = {
'C:\Users\Alvin\Desktop\data_proc\Subj02\RESTING_f002-0004-00001-000001-
'C:\Users\Alvin\Desktop\data_proc\Subj02\RESTING_f002-0004-00002-000002-
'C:\Users\Alvin\Desktop\data_proc\Subj02\RESTING_f002-0004-00003-000003-
'C:\Users\Alvin\Desktop\data_proc\Subj02\RESTING_f002-0004-00004-000004-
'C:\Users\Alvin\Desktop\data_proc\Subj02\RESTING_f002-0004-00005-000005-
'C:\Users\Alvin\Desktop\data_proc\Subj02\RESTING_f002-0004-00006-000006-
'C:\Users\Alvin\Desktop\data_proc\Subj02\RESTING_f002-0004-00007-000007-
'C:\Users\Alvin\Desktop\data_proc\Subj02\RESTING_f002-0004-00008-000008-

```

# Independent Component Analysis



A cocktail-party problem



$$\begin{aligned} x_1(t) &= 0.7s_1(t) + 0.2s_2(t) + 0.1s_3(t) \\ x_2(t) &= 0.3s_1(t) + 0.4s_2(t) + 0.3s_3(t) \\ x_3(t) &= 0.1s_1(t) + 0.2s_2(t) + 0.7s_3(t) \end{aligned}$$

$$\begin{bmatrix} x_1(t) \\ x_2(t) \\ \vdots \\ x_m(t) \end{bmatrix} = \mathbf{A} \begin{bmatrix} s_1(t) \\ s_2(t) \\ \vdots \\ s_k(t) \end{bmatrix}$$

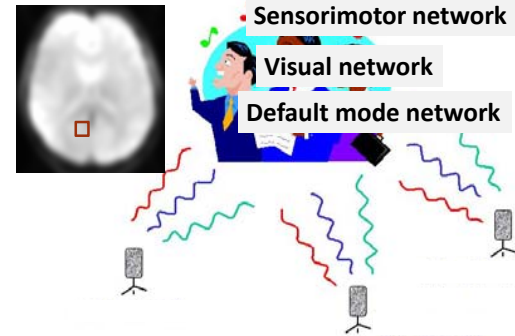
Number of microphones  $\geq$  number of speakers

A is the mixing matrix

# Independent Component Analysis



A cocktail-party problem



$$\begin{aligned} x_1(t) &= 0.7s_1(t) + 0.2s_2(t) + 0.1s_3(t) \\ x_2(t) &= 0.3s_1(t) + 0.4s_2(t) + 0.3s_3(t) \\ x_3(t) &= 0.1s_1(t) + 0.2s_2(t) + 0.7s_3(t) \end{aligned}$$

$$\begin{bmatrix} x_1(t) \\ x_2(t) \\ \vdots \\ x_m(t) \end{bmatrix} = \mathbf{A} \begin{bmatrix} s_1(t) \\ s_2(t) \\ \vdots \\ s_k(t) \end{bmatrix}$$

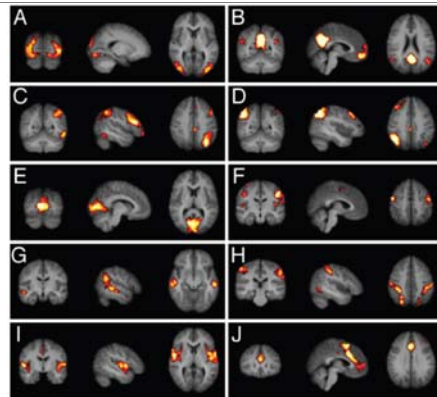
Number of microphones  $\geq$  number of speakers

A is the mixing matrix

# Independent components

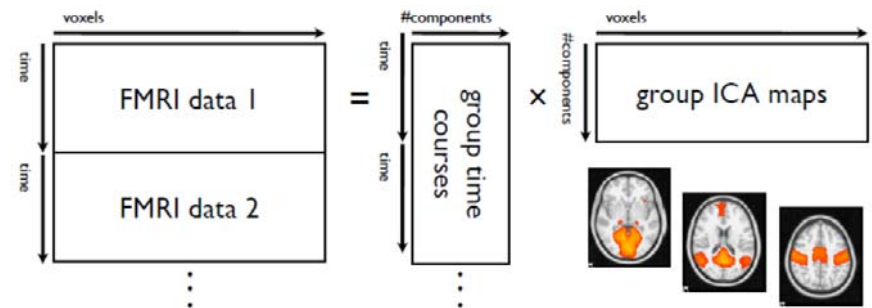


- A. parts of visual cortex
- B. default mode network
- C & D. left and right memory function
- E. visual cortex
- F. sensorimotor cortex
- G. occipitotemporal pathway (ventral stream)
- H. superior parietal cortex
- I. auditory cortex
- J. executive control & working memory



Consistent resting-state networks across healthy subjects. PNAS 2006, 103 37: 13848-13853.

# Group ICA/ Concat-ICA



Group comparison of resting-state FMRI data using multi-subject ICA and dual regression



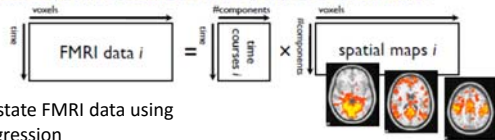
# Dual Regression

③ **Dual Regression:** This is used to identify, within each of the  $N$  individual subjects' fMRI data, spatial maps and associated timecourses corresponding to the multi-subject ICA components. For each individual dataset separately:

(A) use the group-level spatial maps as a set of spatial regressors in a GLM, to find temporal dynamics associated with each group-level map

(B) normalization of these timecourses to unit variance (optional, depending on what question the experimenter wants to ask later)

(C) use these timecourses as a set of temporal regressors in a GLM, to find subject specific maps (still associated with the group-level spatial maps):



Group comparison of resting-state fMRI data using multi-subject ICA and dual regression

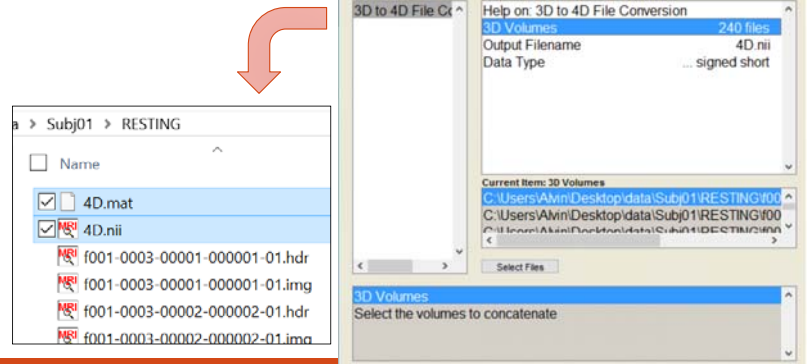
# Q & A

# THE END

[alvin4016@ym.edu.tw](mailto:alvin4016@ym.edu.tw)

# 3D to 4D File Conversion

Batch → SPM → Util → 3D to 4D File Conversion



# Create Your First Batch

