# Neuroimaging data analysis

2nd-level analysis (or group-level inference) in SPM12

#### Sunghyon Kyeong, PhD

Severance Biomedical Science Institute, Yonsei University College of Medicine

#### 실습 데이터 다운로드

Raw data:

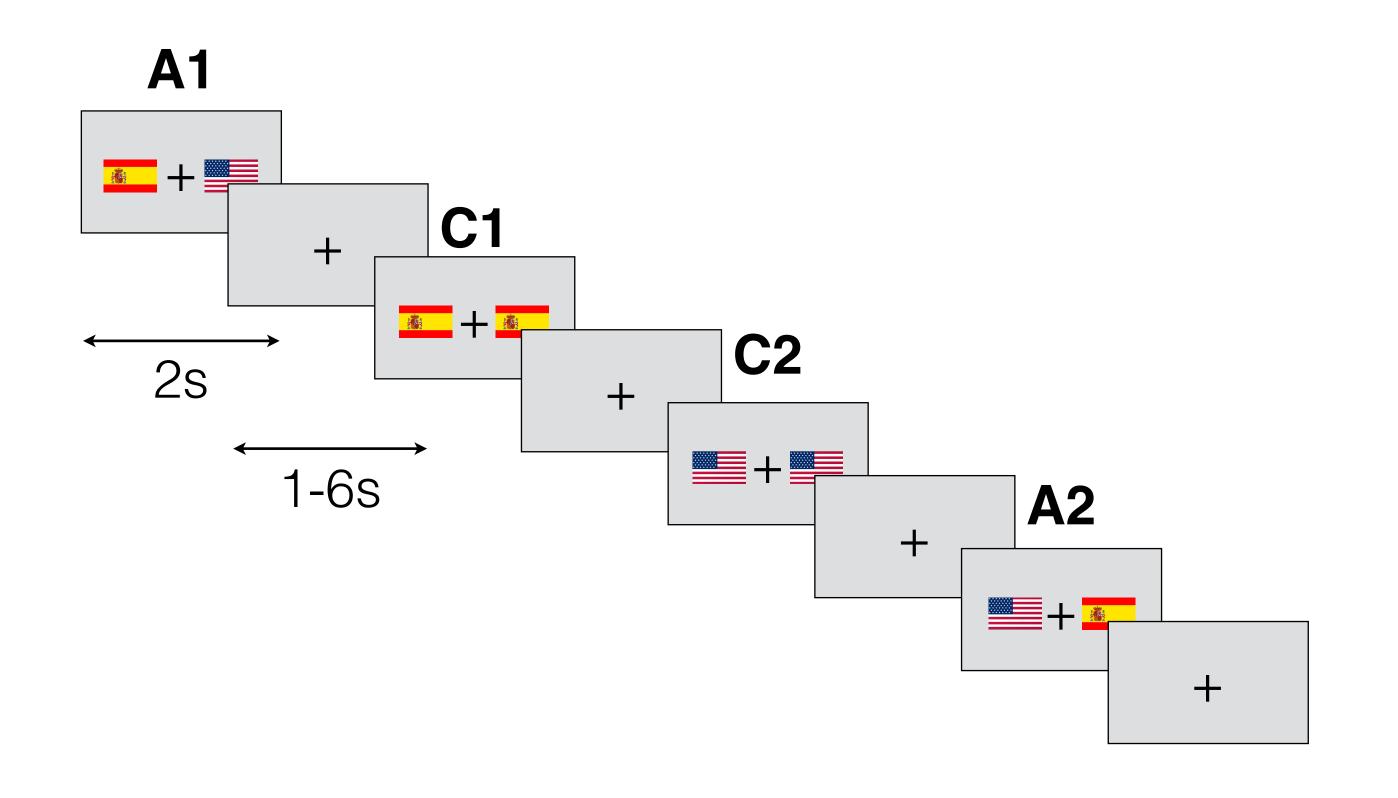
https://openfmri.org/dataset/ds000051/

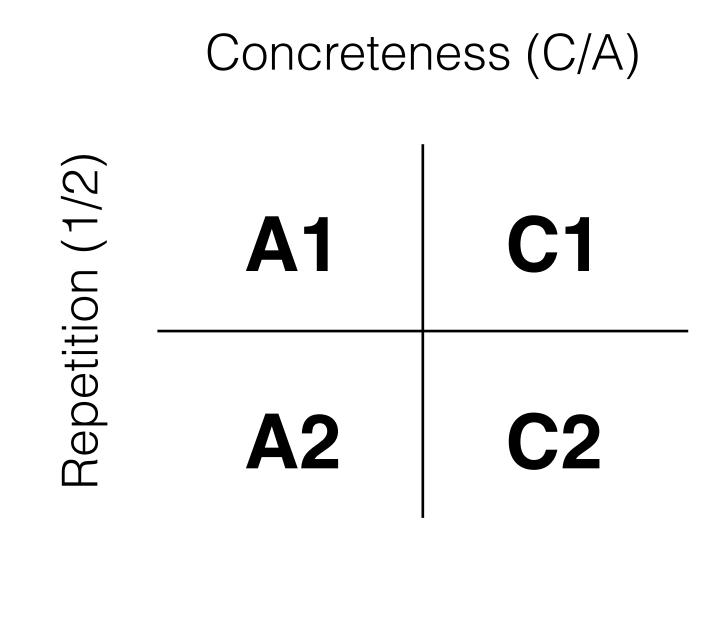
First-level 분석이 완료된 data:

https://drive.google.com/drive/folders/0B6863JbB\_YV-bWFoUE5sbmRvOXc

## 어떤 데이터 일까?

Cross-language repetition priming



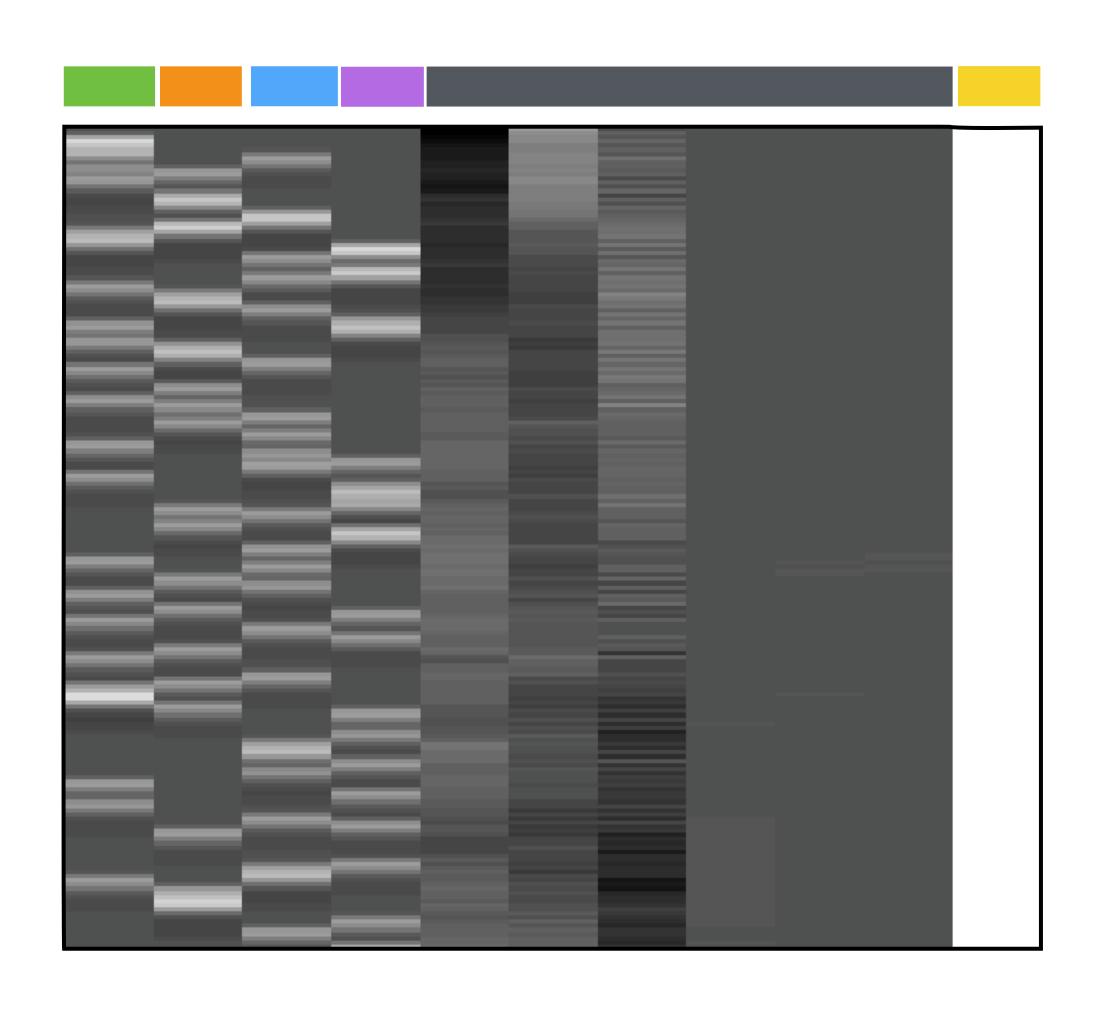


# **Experiment Information**

- (4 x) 2 x 2 factorial event-related fMRI design
- 8 runs (13 subjects)
- (SE, SS, ES, EE) x (abstract vs. concrete) x (1st vs 2nd presentation) of words
- MRI data were acquired using 3T Siemens Allegra system
- 200 Continuous EPI (TE=30ms,TR=2s) per run
- 21 axial slices, 5mm thickness
- 200 mm field of view (64x64 3.125x3.125mm²)

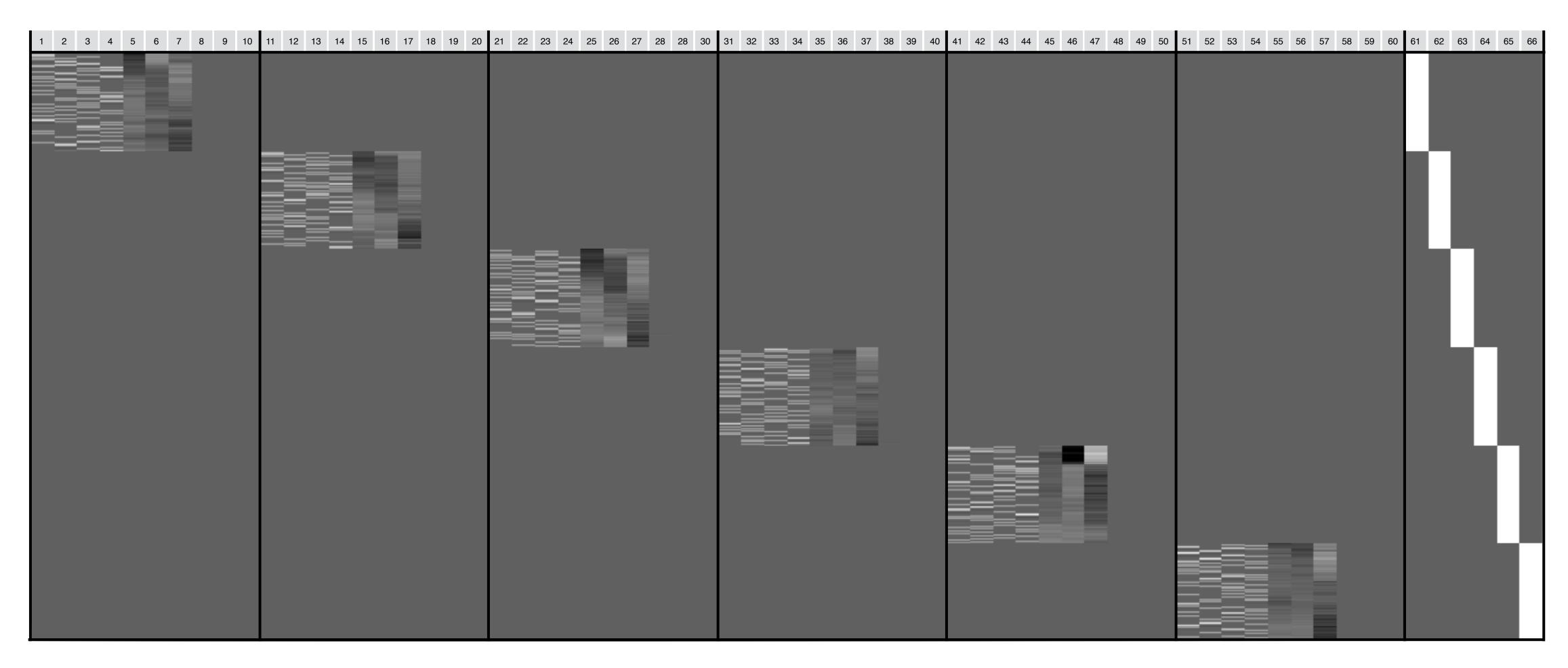
Reference) https://openfmri.org/dataset/ds000051/

#### GLM for Each fMRI Run



- **AN**: Abstract and novel word
- **AR**: Abstract and repeated word
- **CN**: Concrete and novel word
- **CR**: Concrete and repeated word
- Head motion: six rigid head motion parameters
- Constant: a constant term in general linear model

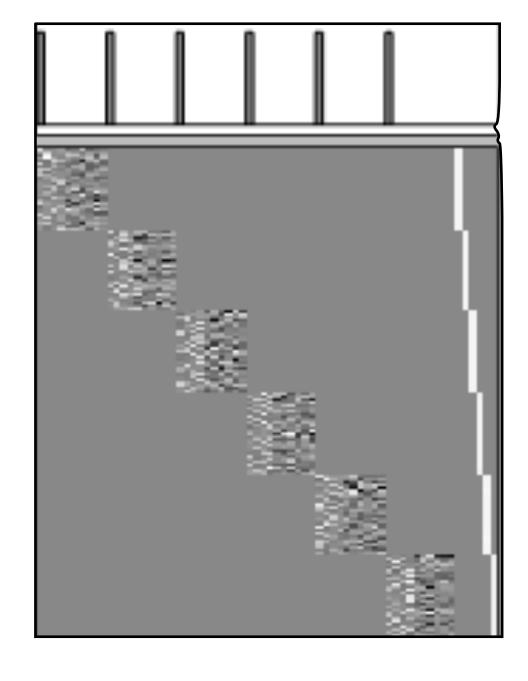
# 1st-level Linear Modeling for 6 Runs



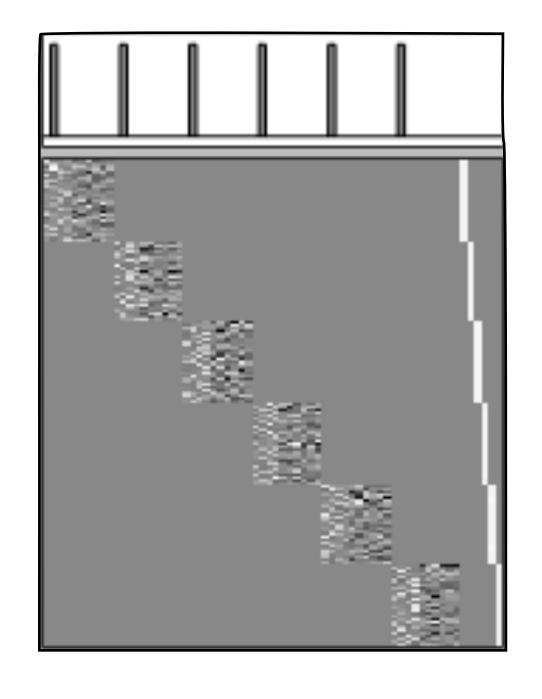
A total of 66 parameters (66 beta images) were estimated.

# Create Contrast Images

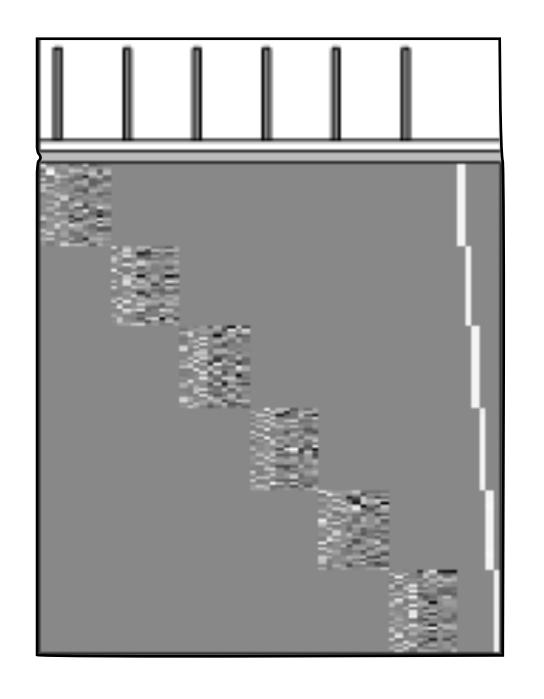
AN 조건 con\_0001.nii



AR 조건 con\_0002.nii



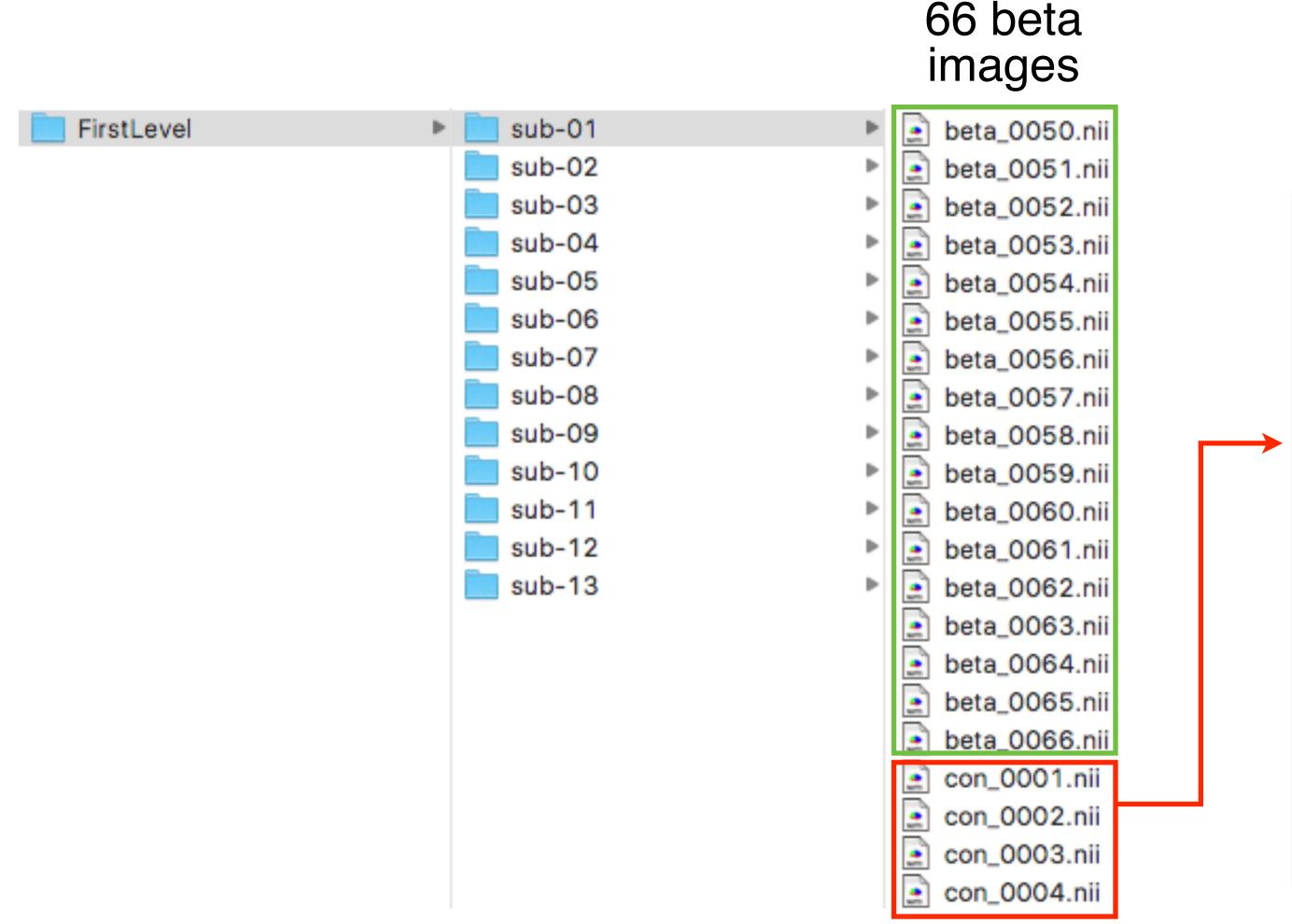
**CN 조건** con\_0003.nii



**CR 조건** con\_0004.nii



#### Contrast Images가 의미하는 바는?



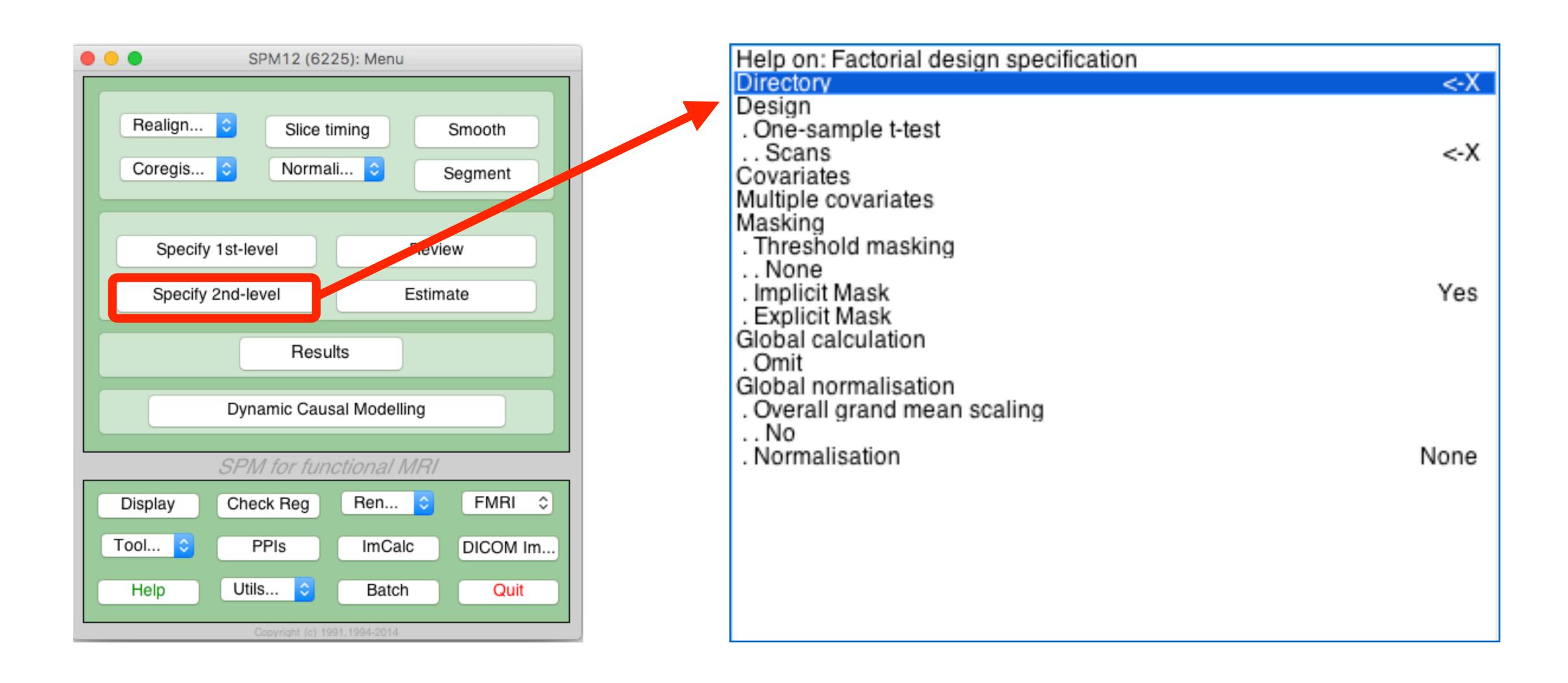
Please note that contrast image is not a average but a summation of interests

$${f con\_0001.nii}$$
  $C_1=\sum_{i=1,11,21,31,41,51}eta_i$  (모든 run의 AN조건 합)  $C_2=\sum_{i=2,12,22,32,42,52}eta_i$  (모든 run의 AR조건 합)  $C_3=\sum_{i=3,13,23,33,43,53}eta_i$  (모든 run의 CN조건 합)  $C_4=\sum_{i=4,14,24,34,44,54}eta_i$  (모든 run의 CR조건 합)

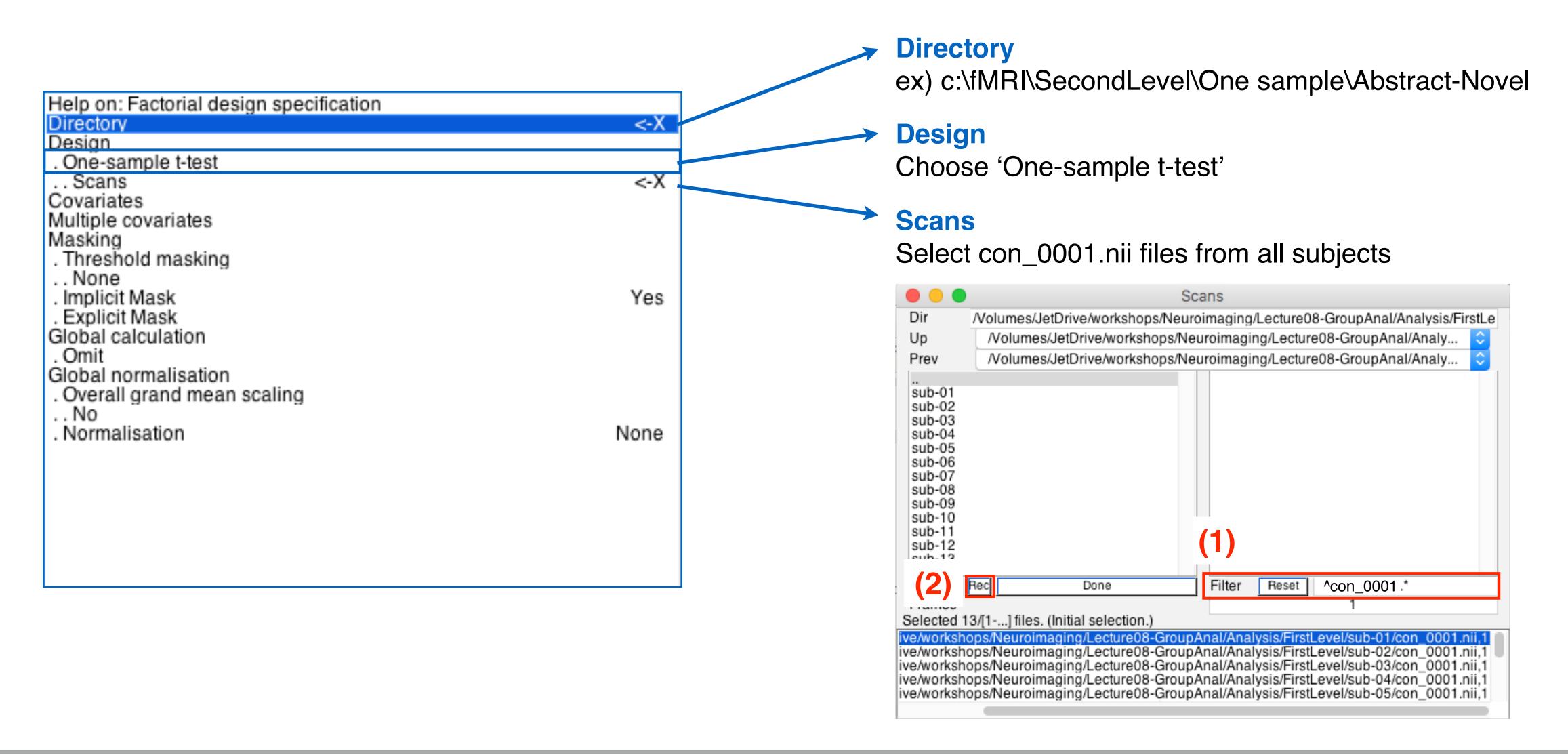
# 그룹 통계 분석 (2nd-level analysis)

- a. one sample t-test
- b. paired sample t-test
- c. full factorial
- d. flexible factorial

## One-sample t-test: Specify model (1/2)

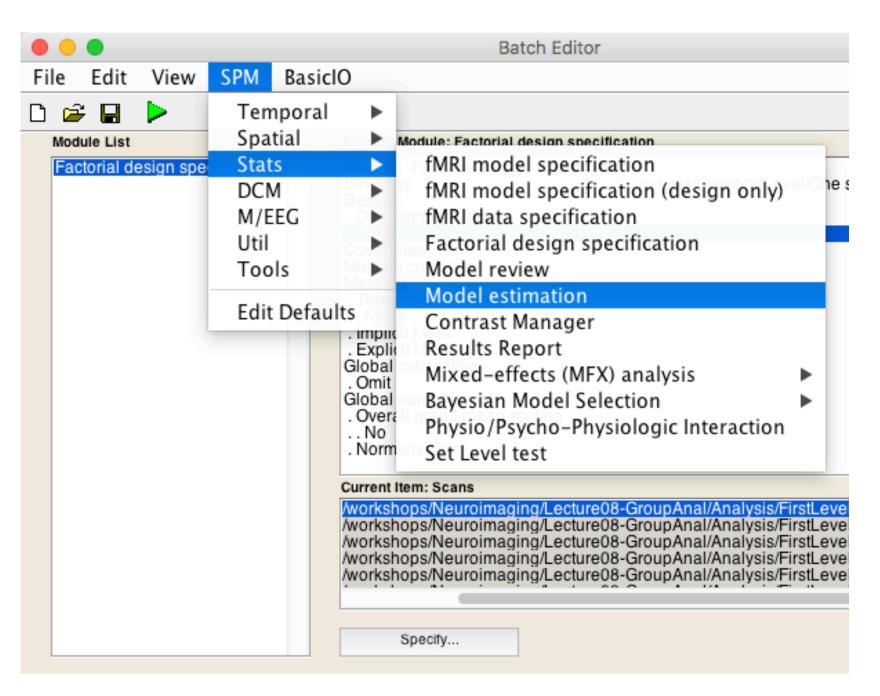


# One-sample t-test: Specify model (2/2)

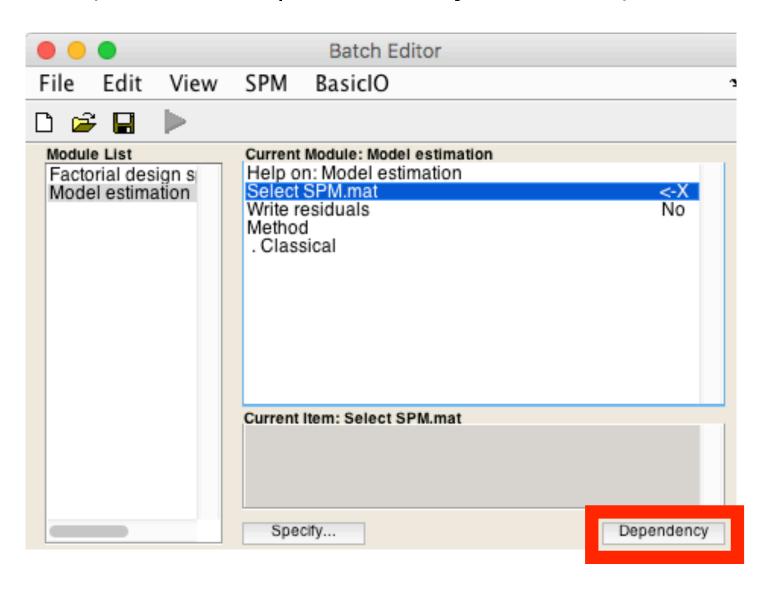


#### One-sample t-test: Estimation

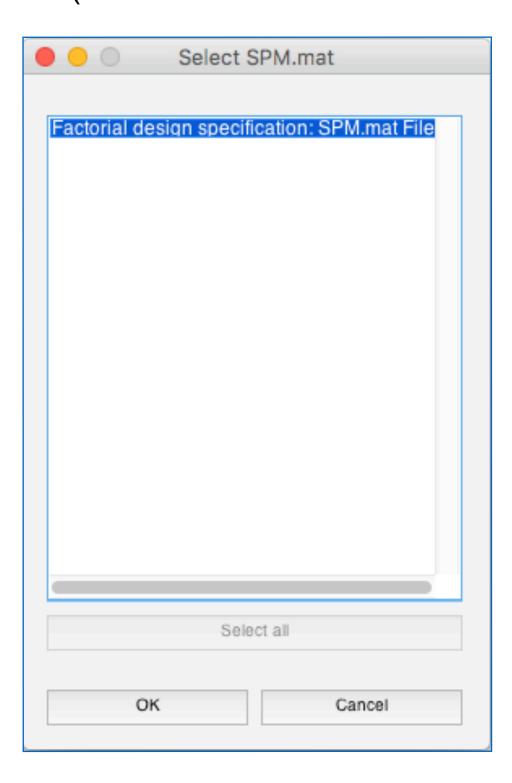
Add 'model estimation' in the module list
 (SPM menu → Stats → Model estimation)



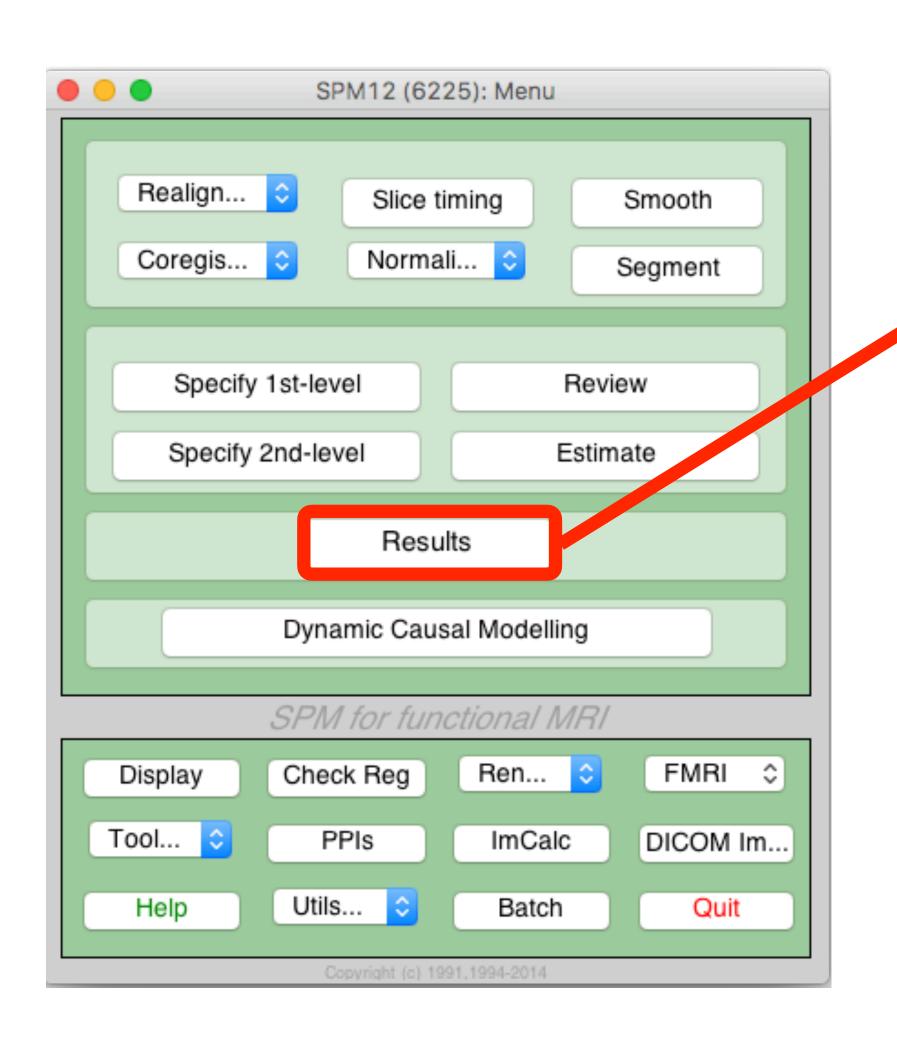
2. Select SPM.mat file (Click 'Dependency' button)



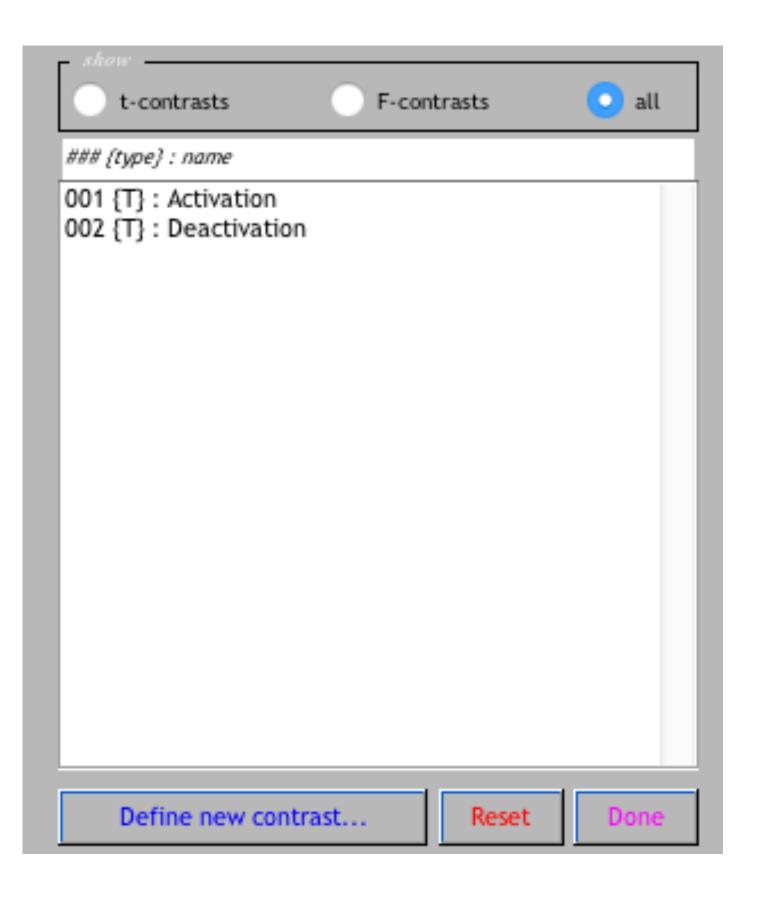
3. Select SPM.mat file (click Run after selection)



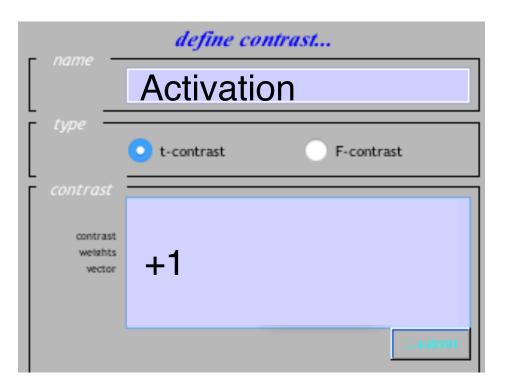
#### One-sample t-test: Contrast



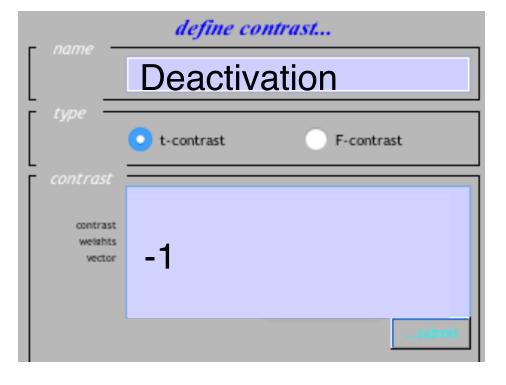
Select one of existing contrasts or Define net contrast



#### Activation-contrast



#### Deactivation-contrast



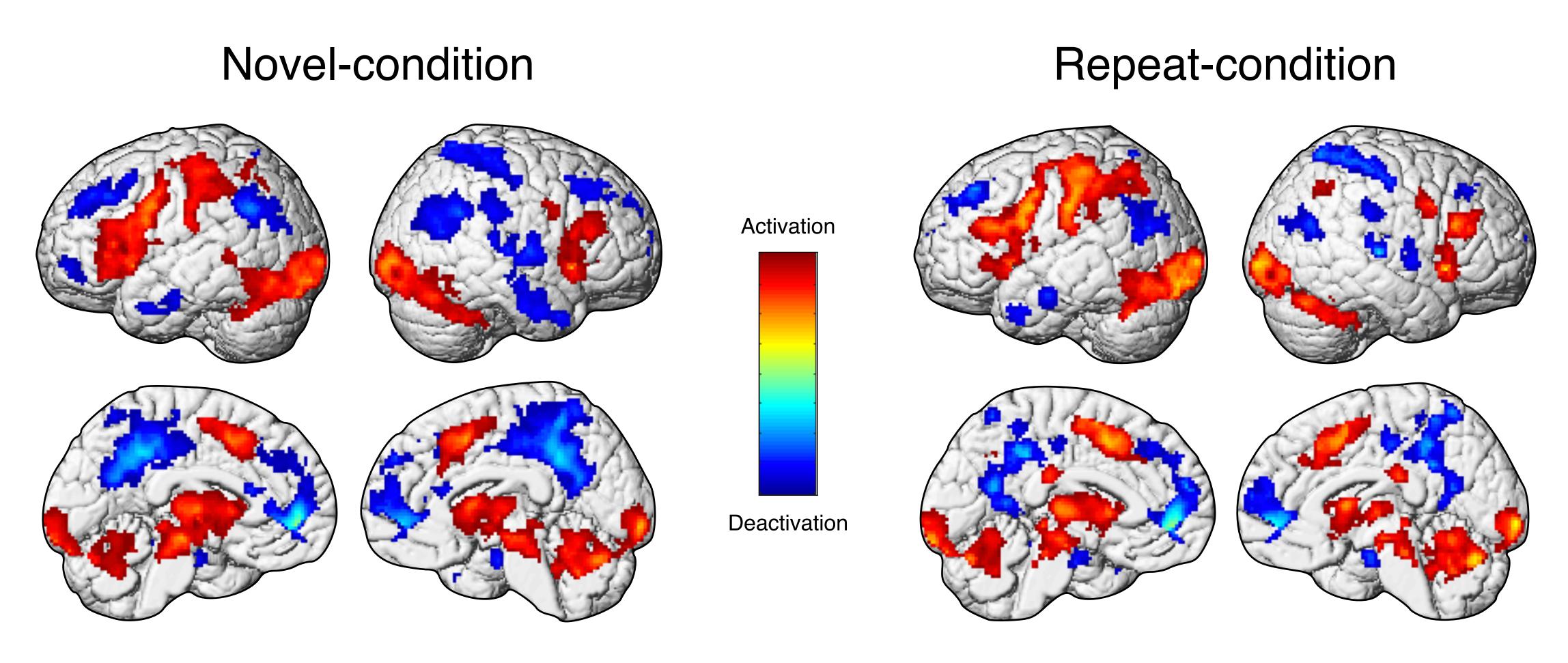
# 같은 방법으로 AN, AR, CN, CR 조건에 대해서 One-sample t-test 분석을 진행하고 그 결과를 확인해 본다.

#### Matlab script for one-sample t-test

```
>> % Directory containing Cross-language data
>> %-----
>> proj path = '/Volumes/JetDrive/workshops/Neuroimaging/Lecture08-GroupAnal';
>> fn xls = fullfile(proj path, 'subjlist.xlsx');
>>
>> T = readtable(fn xls);
>> subjlist = T.subjname;
>> nsubj = length(subjlist);
>>
>> dir name = 'Abstract-Novel';
>> con name = 'con 0001.nii';
>>
>>
>> % Specify Model
>> clear matlabbatch;
>> out_dir = fullfile(proj_path, 'Analysis', 'SecondLevel', 'One sample', dir_name); mkdir(out_dir);
>> confiles = cell(0);
>> for c=1:nsubj,
      confiles{c} = fullfile( proj_path, 'Analysis', 'FirstLevel', subjlist{c}, con_name);
>> end
>> matlabbatch{1}.spm.stats.factorial design.dir = {out dir};
>> matlabbatch{1}.spm.stats.factorial_design.des.t1.scans = [confiles'];
>> matlabbatch{1}.spm.stats.factorial design.masking.im = 1;
```

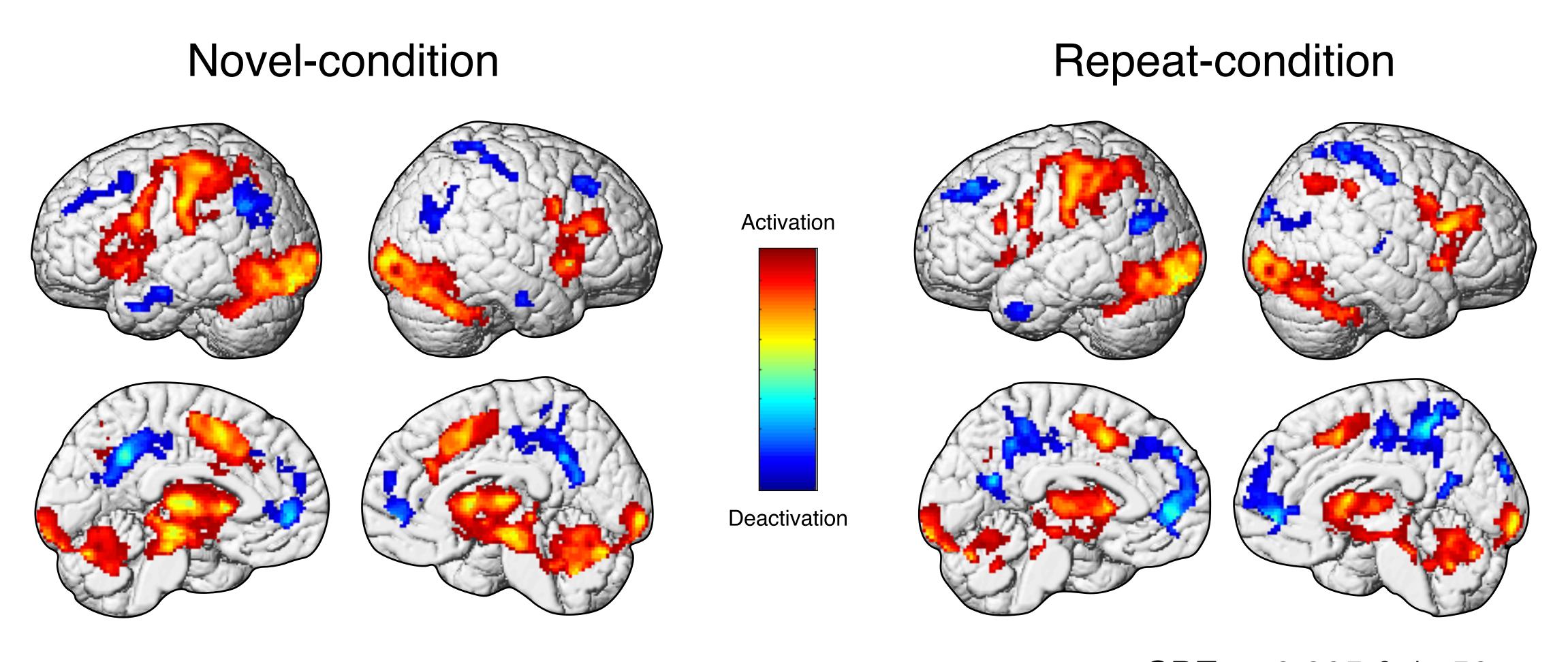
```
>> % Parameter Estimation
>> SPM mat = fullfile(out dir, 'SPM.mat');
>> matlabbatch{2}.spm.stats.fmri est.spmmat = {SPM mat};
>> matlabbatch{2}.spm.stats.fmri_est.write_residuals = 0;
>> matlabbatch{2}.spm.stats.fmri_est.method.Classical = 1;
>>
>>
>> % Create Contrasts
>> matlabbatch{3}.spm.stats.con.spmmat = {SPM mat};
>> matlabbatch{3}.spm.stats.con.consess{1}.tcon.name = 'Activation';
>> matlabbatch{3}.spm.stats.con.consess{1}.tcon.weights = 1;
>> matlabbatch{3}.spm.stats.con.consess{1}.tcon.sessrep = 'none';
>> matlabbatch{3}.spm.stats.con.consess{2}.tcon.name = 'Deactivation';
>> matlabbatch{3}.spm.stats.con.consess{2}.tcon.weights = -1;
>> matlabbatch{3}.spm.stats.con.consess{2}.tcon.sessrep = 'none';
>> matlabbatch{3}.spm.stats.con.delete = 1;
>> % spm jobman('interactive', matlabbatch);
>> spm jobman('run', matlabbatch);
```

#### Abstract word condition



CDT *p*<0.005 & *k*=50

#### Concrete word condition

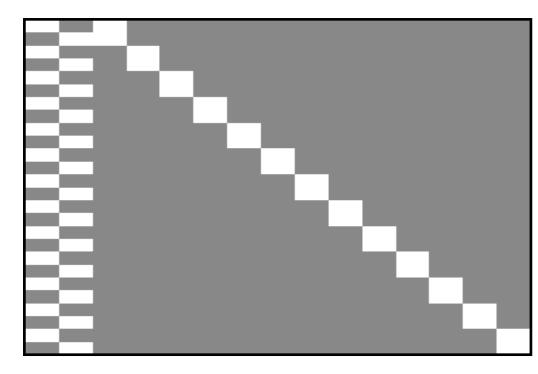


#### 무엇을 알 수 있을까?

- One-sample t-test의 결과로 각각의 실험 조건에서 뇌가 활성화 되거나 비활성화 되는 영역을 알 수 있다.
- 하지만, 실험 조건간에 어느 영역이 더욱 활성화 되고, 어느 영역이 덜 활성화 되는지 여부는 알 수 없다. 하지만, 각각의 실험 조건에 관여하는 뇌의 영역을 발견하는 데에는 효과적인 통계분석 방법이다.

# 그룹 통계 분석 (2nd-level analysis)

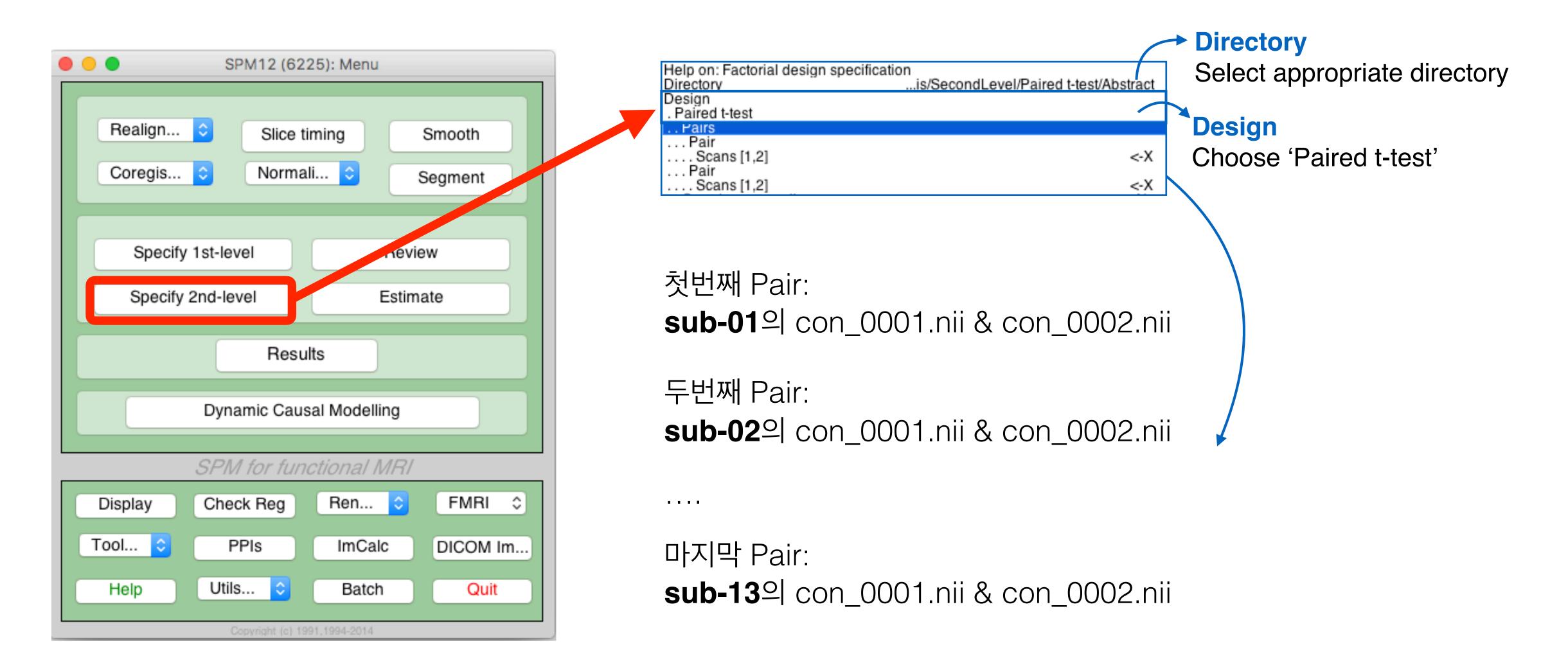
- a. one sample t-test
- b. paired sample t-test
- c. full factorial
- d. flexible factorial



Design matrix of paired sample model

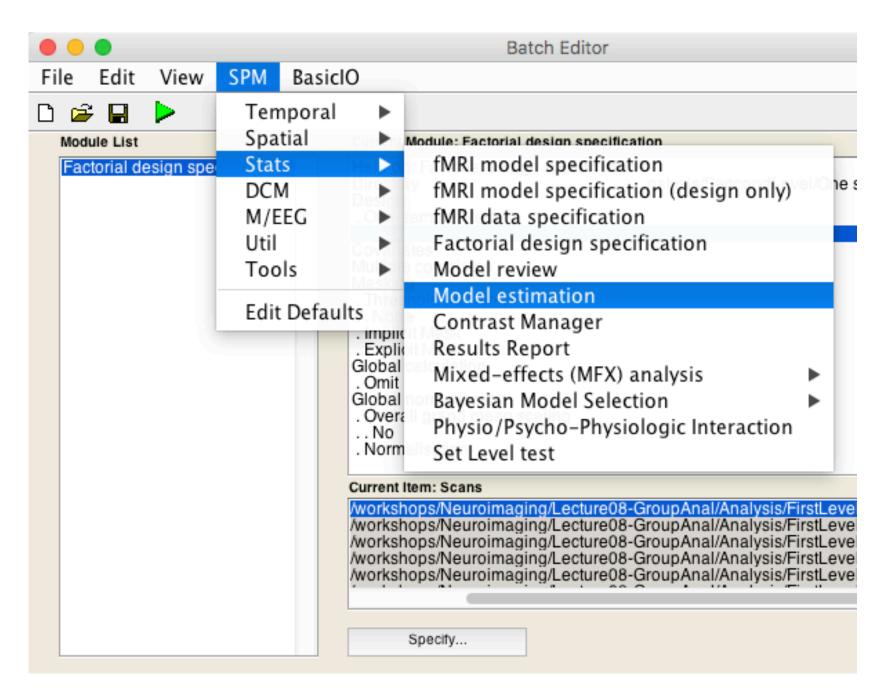
## Paired sample t-test: Specify model

Abstract word 단어가 처음 제시되었을 때와 반복되어 제시되었을 때의 뇌반응 차이

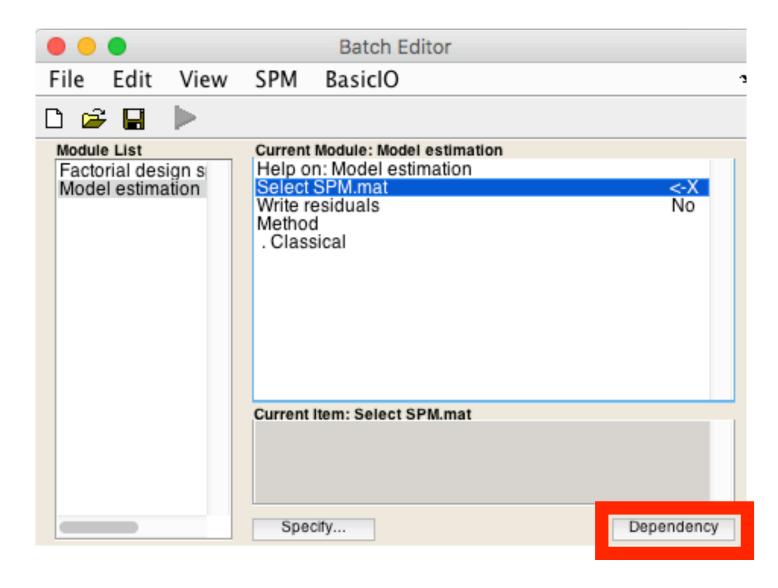


#### Paired sample t-test: Estimation

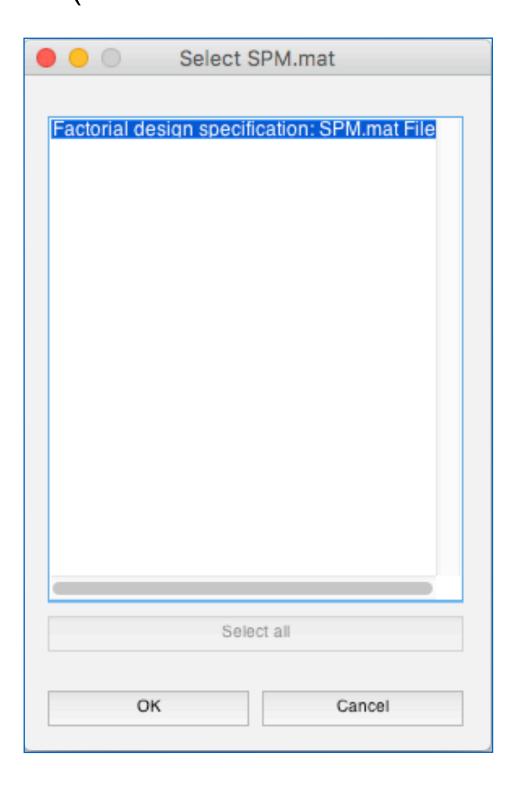
Add 'model estimation' in the module list
 (SPM menu → Stats → Model estimation)



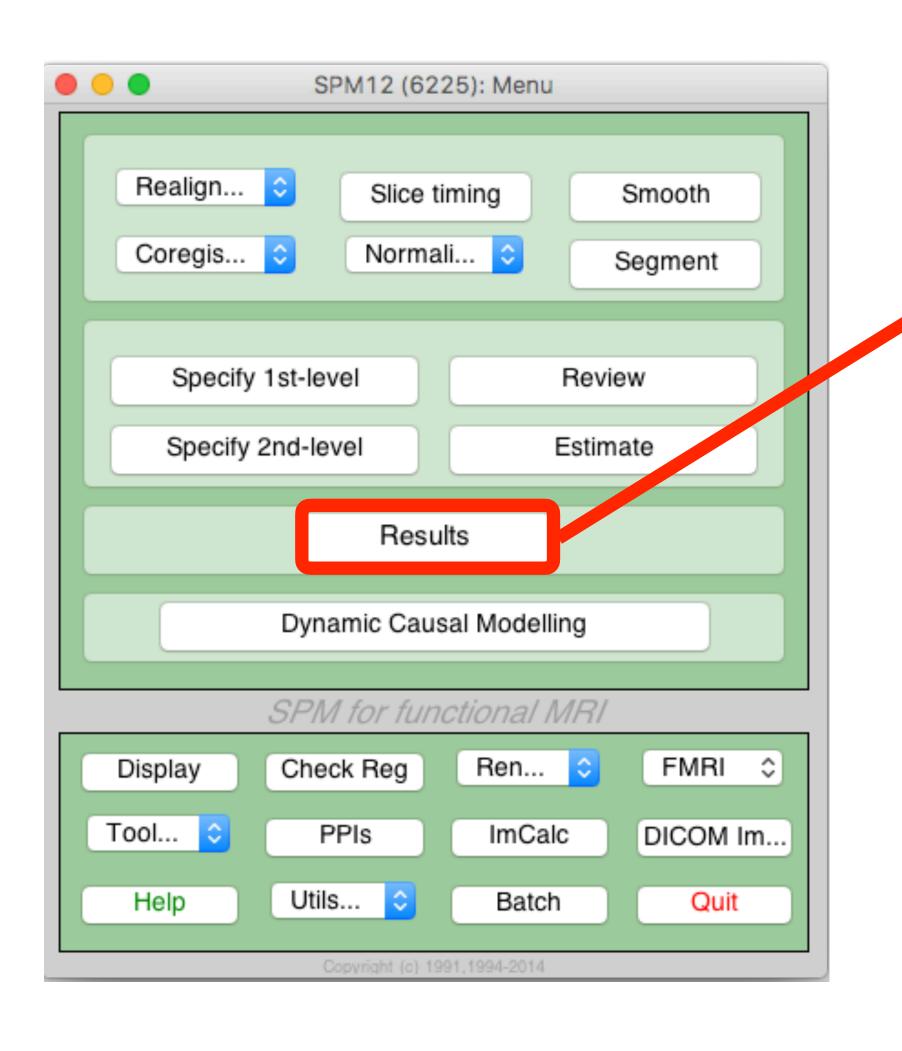
2. Select SPM.mat file (Click 'Dependency' button)



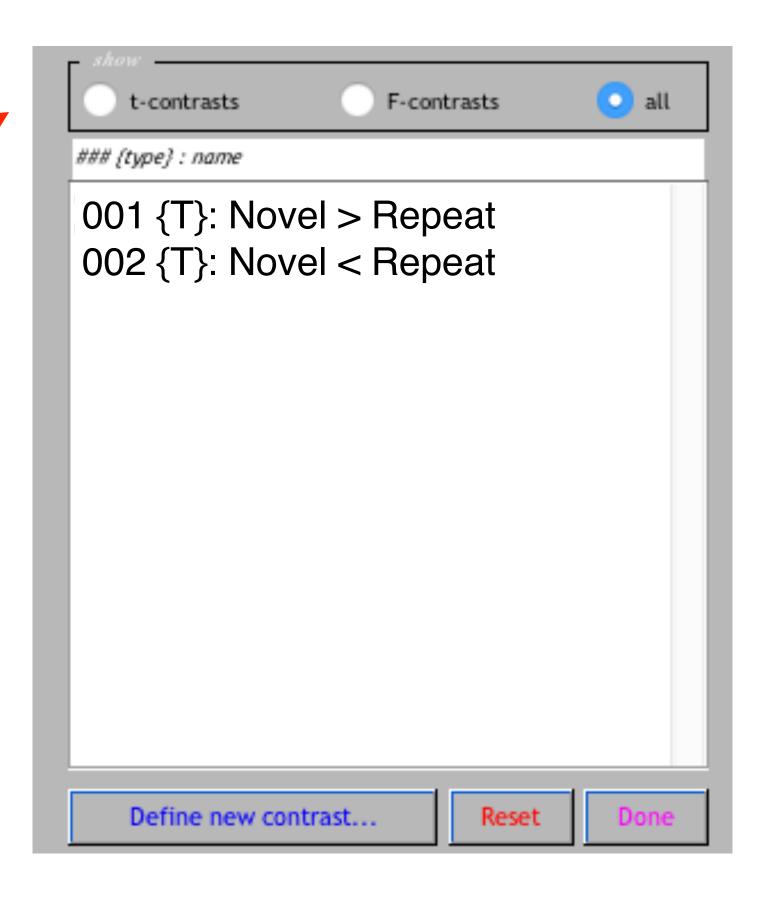
3. Select SPM.mat file (click > Run after selection)



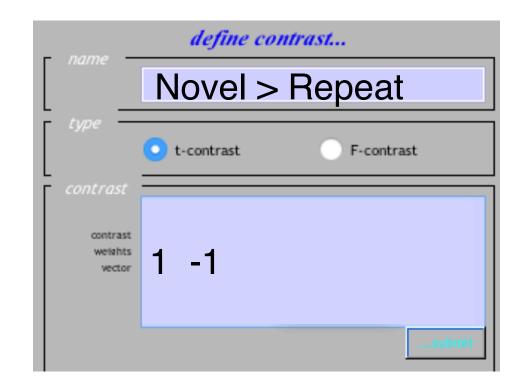
#### Paired sample t-test: Contrast



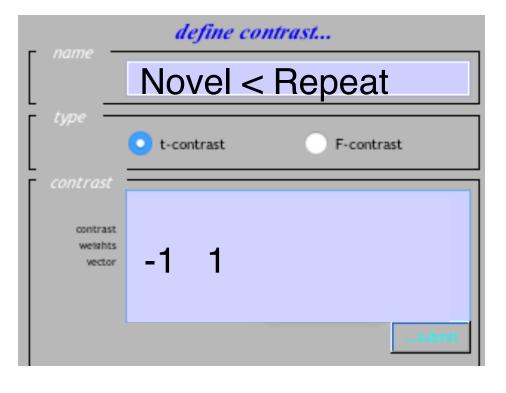
Select one of existing contrasts or Define net contrast



#### Contrast of Novel-Repeat



#### Contrast of Repeat-Novel



#### Matlab script for paired sample t-test

```
>> % Directory containing Cross-language data
>> %------
>> proj_path = '/Volumes/JetDrive/workshops/Neuroimaging/Lecture08-GroupAnal';
>> fn_xls = fullfile(proj_path,'subjlist.xlsx');
>>
>> T = readtable(fn xls);
>> subjlist = T.subjname;
>> nsubj = length(subjlist);
>>
>> dir_name = 'Concrete';
>> con novel = 'con 0003.nii';
>> con repeat = 'con 0004.nii';
>>
>>
>> % Specify Model
>> clear matlabbatch;
>> out dir = fullfile(proj path, 'Analysis', 'SecondLevel', 'Paired t-test', dir name);
  mkdir(out dir);
>> matlabbatch{1}.spm.stats.factorial design.dir = {out dir};
>> for c=1:nsubj,
      paired_confiles = cell(0);
      paired confiles{1} =
  fullfile( proj_path, 'Analysis', 'FirstLevel', subjlist{c}, con_novel);
      paired confiles{2} =
  fullfile( proj path, 'Analysis', 'FirstLevel', subjlist(c), con repeat);
      matlabbatch{1}.spm.stats.factorial_design.des.pt.pair(c).scans =
  [paired_confiles'];
>> end
```

```
>> % Parameter Estimation
>> SPM_mat = fullfile(out_dir,'SPM.mat');
>> matlabbatch{2}.spm.stats.fmri_est.spmmat = {SPM_mat};
>> matlabbatch{2}.spm.stats.fmri est.write residuals = 0;
>> matlabbatch{2}.spm.stats.fmri_est.method.Classical = 1;
>>
>>
>> % Create Contrasts
>> matlabbatch{3}.spm.stats.con.spmmat = {SPM mat};
>> matlabbatch{3}.spm.stats.con.consess{1}.tcon.name = 'Novel > Repeat';
>> matlabbatch{3}.spm.stats.con.consess{1}.tcon.weights = [1 -1];
>> matlabbatch{3}.spm.stats.con.consess{1}.tcon.sessrep = 'none';
>> matlabbatch{3}.spm.stats.con.consess{2}.tcon.name = 'Novel < Repeat';
>> matlabbatch{3}.spm.stats.con.consess{2}.tcon.weights = [-1 1];
>> matlabbatch{3}.spm.stats.con.consess{2}.tcon.sessrep = 'none';
>> matlabbatch{3}.spm.stats.con.delete = 1;
>> % spm_jobman('interactive', matlabbatch);
>> spm jobman('run', matlabbatch);
```

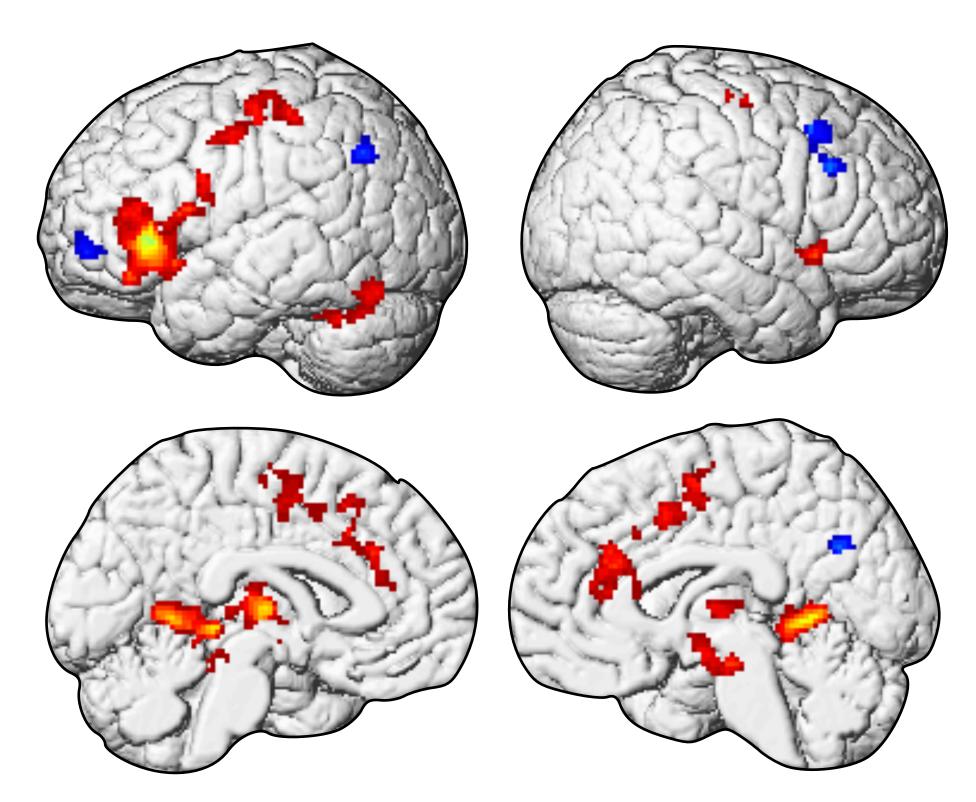
## Paired sample t-test: Mapping

Novel > Repeat

Novel < Repeat

Abstract word condition (Novel-Repeat)

Concrete word condition (Novel-Repeat)



CDT *p*<0.005 & *k*=50

#### 무엇을 알 수 있을까?

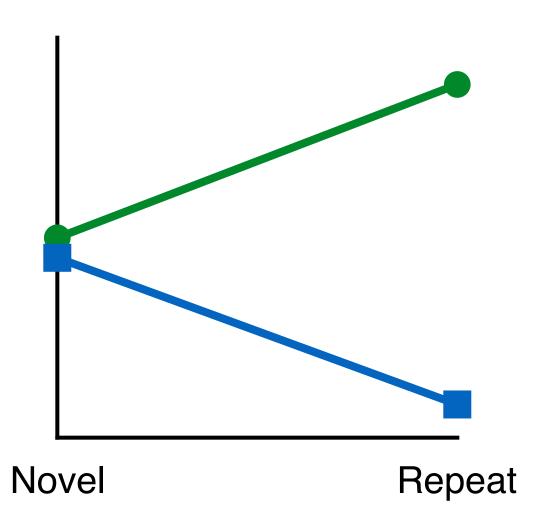
Paired sample t-test의 결과로 Pair로 생각할 수 있는 두개의 조건들
 간에 통계적으로 차이가 있는 영역들을 알 수 있다.

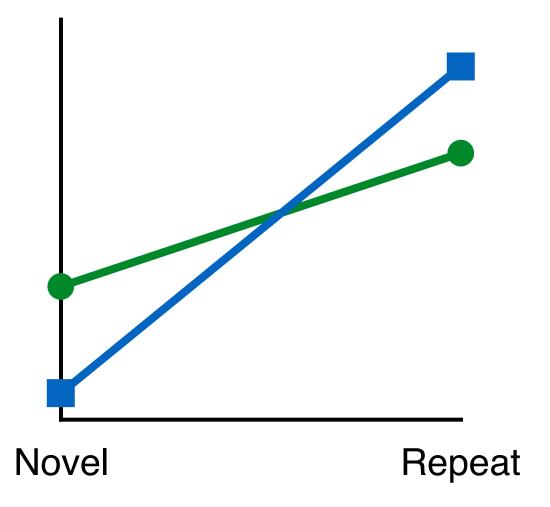
• 하지만, 상호작용 효과는 알 수 없다.

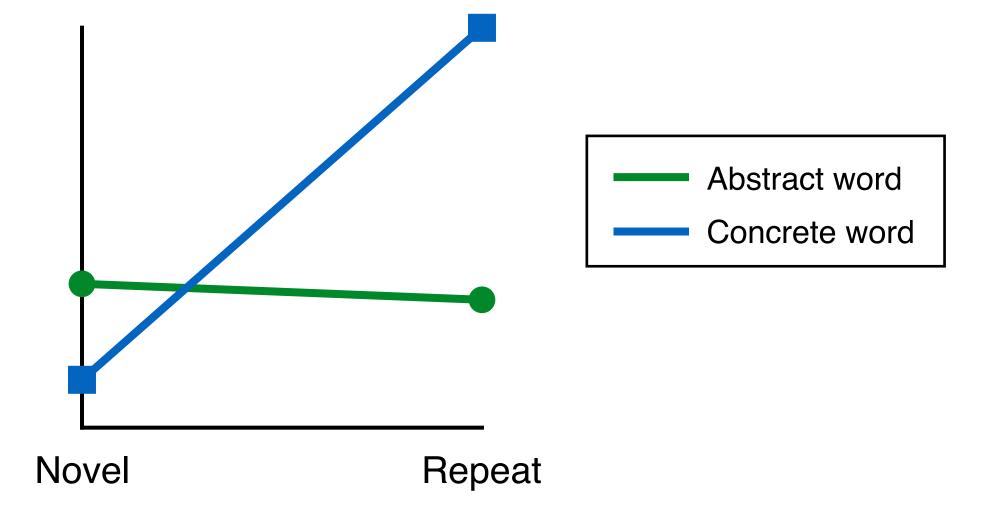
# 상호작용 (Interaction)

Interaction, Main effect of word condition Interaction, Main effect of repetition

Interaction, Main effect of word condition Main effect of repetition

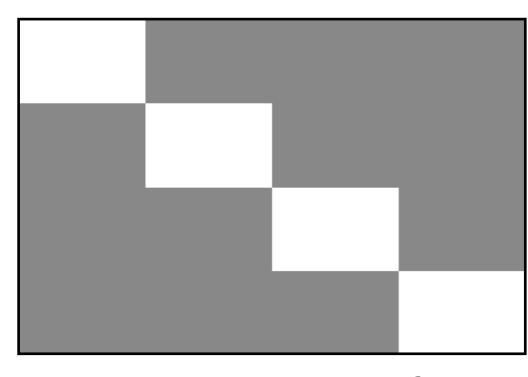






# 그룹 통계 분석 (2nd-level analysis)

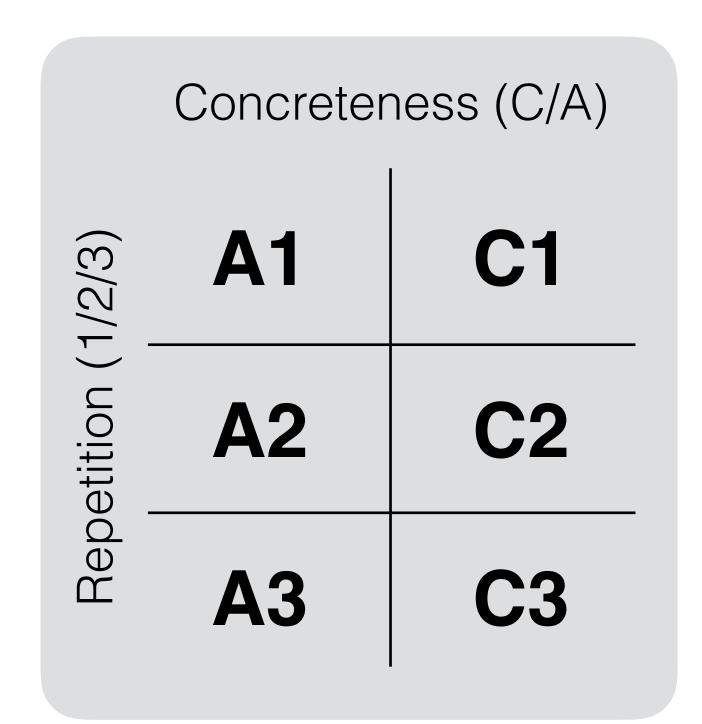
- a. one sample t-test
- b. paired sample t-test
- c. full factorial
- d. flexible factorial



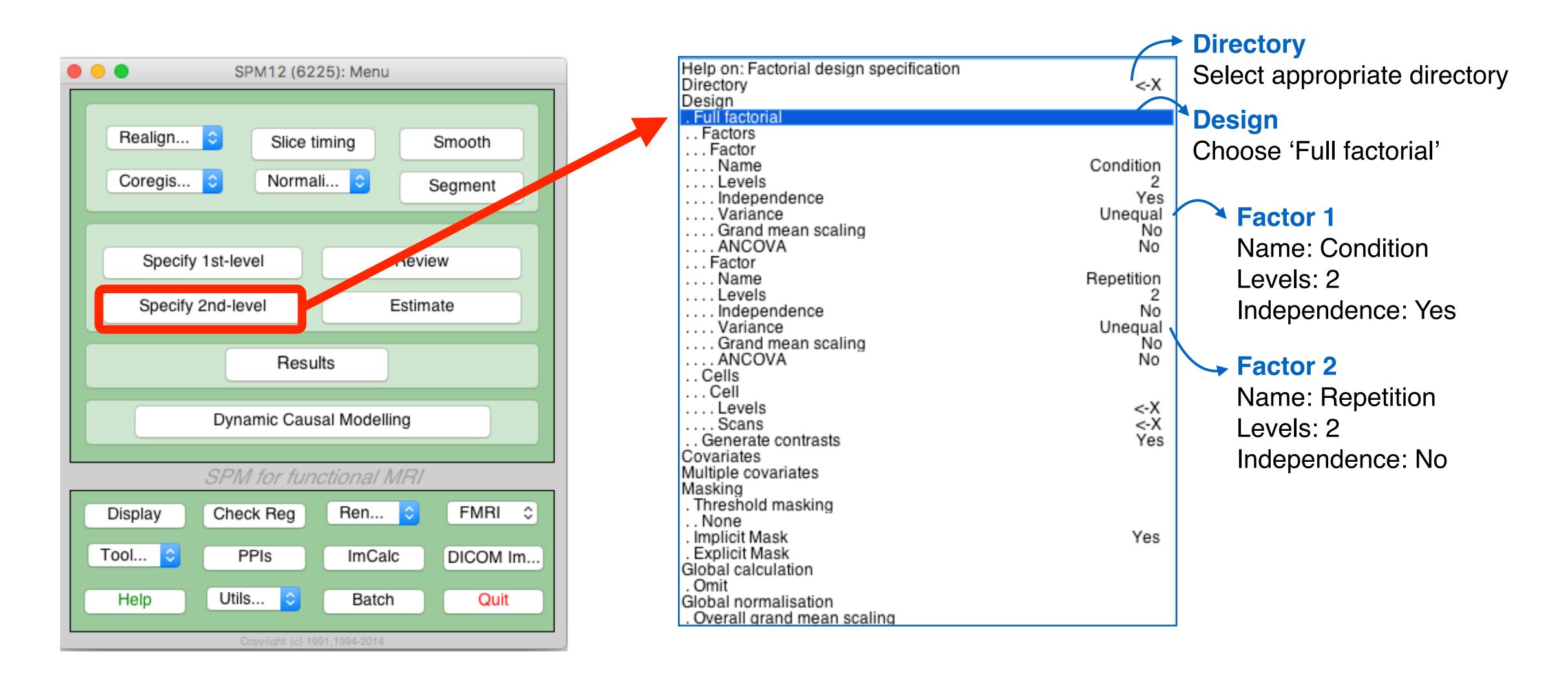
Design matrix of full factorial model

#### Factorial design

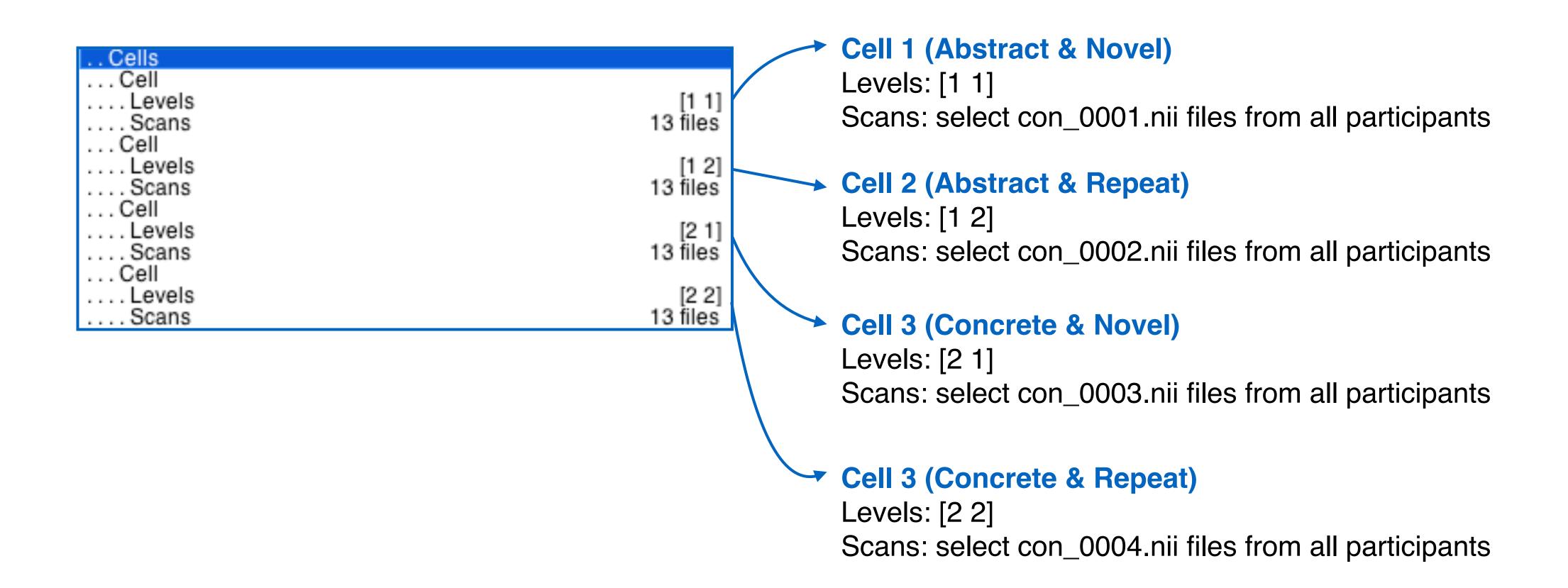
- If you have a factorial design then SPM can automatically generate the contrasts necessary to test for the main effects and interactions.
- This includes the F-contrasts necessary to test for these effects at the within-subject level (first level) and the simple contrasts necessary to generate the contrast images for a between-subject (second-level) analysis.
- What is a factor? For example, if you have 2-by-3
  design in your contingency table, you have two factors.
   One has 2-levels and another has 3-levels.



# Full factorial: Specify model (1/2)

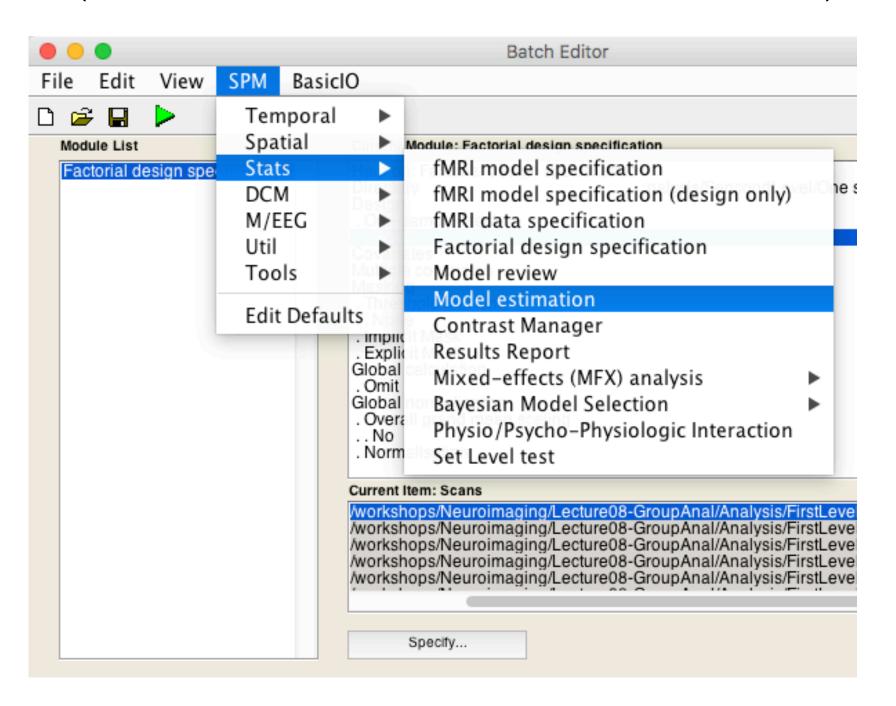


## Full factorial: Specify model (2/2)

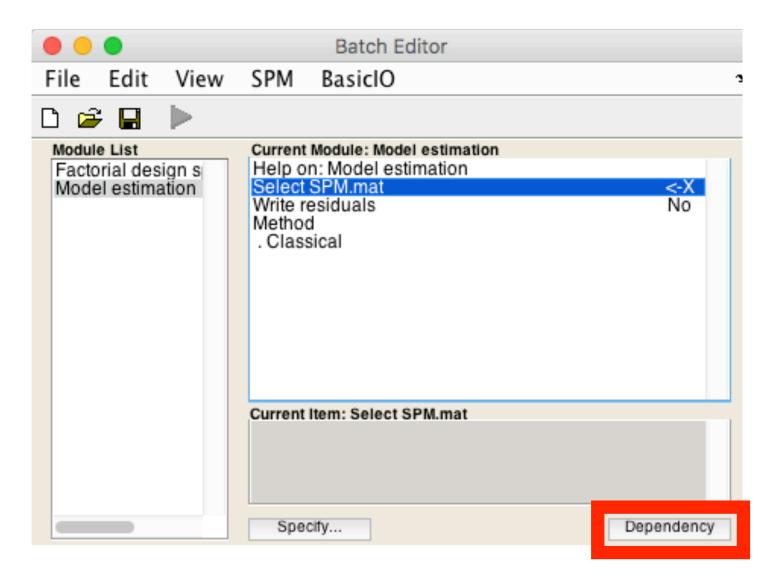


#### Full factorial: Estimation

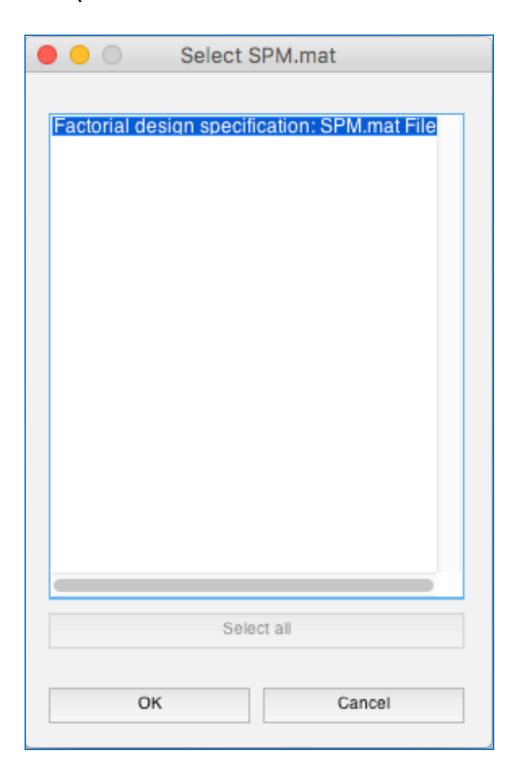
1. Add 'model estimation' in the module list (SPM menu → Stats → Model estimation)



2. Select SPM.mat file (Click 'Dependency' button)



3. Select SPM.mat file (click ▶ Run after selection)



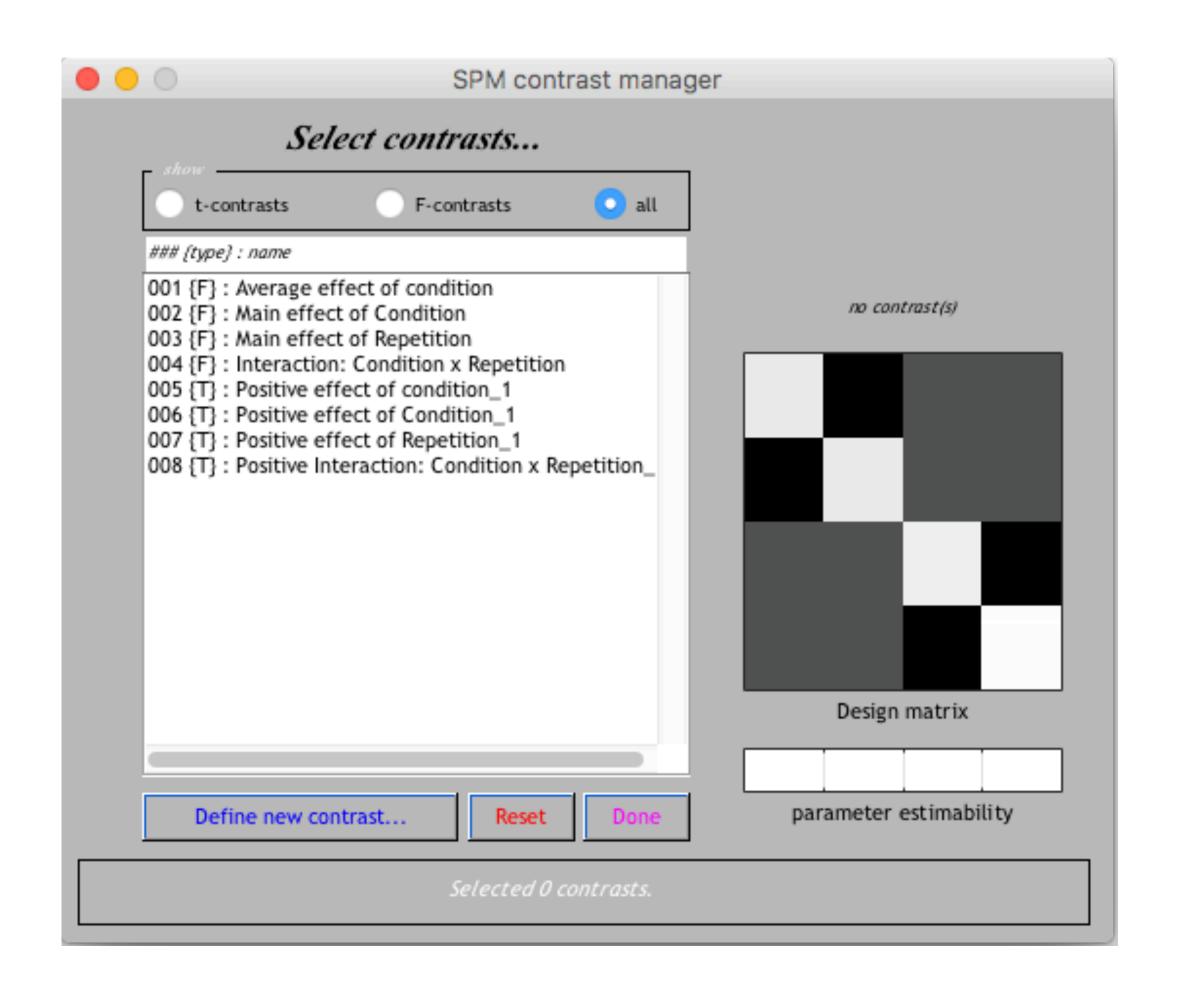
Full factorial design에서는 Estimation을 수행하고 나면, 자동으로 contrasts가 생성됨

## Matlab script for full factorial analysis

```
>> % Directory containing Cross-language data
>> %-----
>> proj path = '/Volumes/JetDrive/workshops/Neuroimaging/Lecture08-GroupAnal';
>> fn_xls = fullfile(proj_path,'subjlist.xlsx');
>>
>> T = readtable(fn xls);
>> subjlist = T.subjname;
>> nsubj = length(subjlist);
>>
>>
>> % Specify Model
>> %-----
>> clear matlabbatch;
>> out dir = fullfile(proj path, 'Analysis', 'SecondLevel', 'Factorial'); mkdir(out dir);
>> confiles = cell(0);
>> con2files = cell(0);
>> con3files = cell(0);
>> con4files = cell(0);
>>
>> for c=1:nsubj,
      con1files{c} = fullfile( proj_path, 'Analysis', 'FirstLevel', subjlist{c}, 'con_0001.nii');
>>
      con2files{c} = fullfile( proj_path, 'Analysis', 'FirstLevel', subjlist{c}, 'con_0002.nii');
>>
      con3files{c} = fullfile( proj_path, 'Analysis', 'FirstLevel', subjlist{c}, 'con_0003.nii');
>>
      con4files{c} = fullfile( proj_path, 'Analysis', 'FirstLevel', subjlist{c}, 'con_0004.nii');
>>
>> end
>>
```

```
>> matlabbatch{1}.spm.stats.factorial design.dir = {out dir};
>> matlabbatch{1}.spm.stats.factorial design.des.fd.fact(1).name = 'Condition';
>> matlabbatch{1}.spm.stats.factorial design.des.fd.fact(1).levels = 2;
>> matlabbatch{1}.spm.stats.factorial design.des.fd.fact(1).dept = 0;
>> matlabbatch{1}.spm.stats.factorial_design.des.fd.fact(1).variance = 1;
>> matlabbatch{1}.spm.stats.factorial design.des.fd.fact(1).gmsca = 0;
>> matlabbatch{1}.spm.stats.factorial design.des.fd.fact(1).ancova = 0;
>> matlabbatch{1}.spm.stats.factorial_design.des.fd.fact(2).name = 'Repetition';
>> matlabbatch{1}.spm.stats.factorial design.des.fd.fact(2).levels = 2;
>> matlabbatch{1}.spm.stats.factorial_design.des.fd.fact(2).dept = 1;
>> matlabbatch{1}.spm.stats.factorial_design.des.fd.fact(2).variance = 1;
>> matlabbatch{1}.spm.stats.factorial design.des.fd.fact(2).gmsca = 0;
>> matlabbatch{1}.spm.stats.factorial_design.des.fd.fact(2).ancova = 0;
>> matlabbatch{1}.spm.stats.factorial design.des.fd.icell(1).levels = [1 1];
>> matlabbatch{1}.spm.stats.factorial design.des.fd.icell(1).scans = [con1files'];
>> matlabbatch{1}.spm.stats.factorial design.des.fd.icell(2).levels = [1 2];
>> matlabbatch{1}.spm.stats.factorial_design.des.fd.icell(2).scans = [con2files'];
>> matlabbatch{1}.spm.stats.factorial_design.des.fd.icell(3).levels = [2 1];
>> matlabbatch{1}.spm.stats.factorial design.des.fd.icell(3).scans = [con3files'];
>> matlabbatch{1}.spm.stats.factorial_design.des.fd.icell(4).levels = [2 2];
>> matlabbatch{1}.spm.stats.factorial design.des.fd.icell(4).scans = [con4files'];
>> matlabbatch{1}.spm.stats.factorial design.des.fd.contrasts = 1;
>>
>>
>> % Parameter Estimation
>> SPM mat = fullfile(out dir, 'SPM.mat');
>> matlabbatch{2}.spm.stats.fmri est.spmmat = {SPM mat};
>> matlabbatch{2}.spm.stats.fmri est.write residuals = 0;
>> matlabbatch{2}.spm.stats.fmri est.method.Classical = 1;
>> % spm jobman('interactive', matlabbatch);
>> spm jobman('run', matlabbatch);
```

#### Full factorial: Results

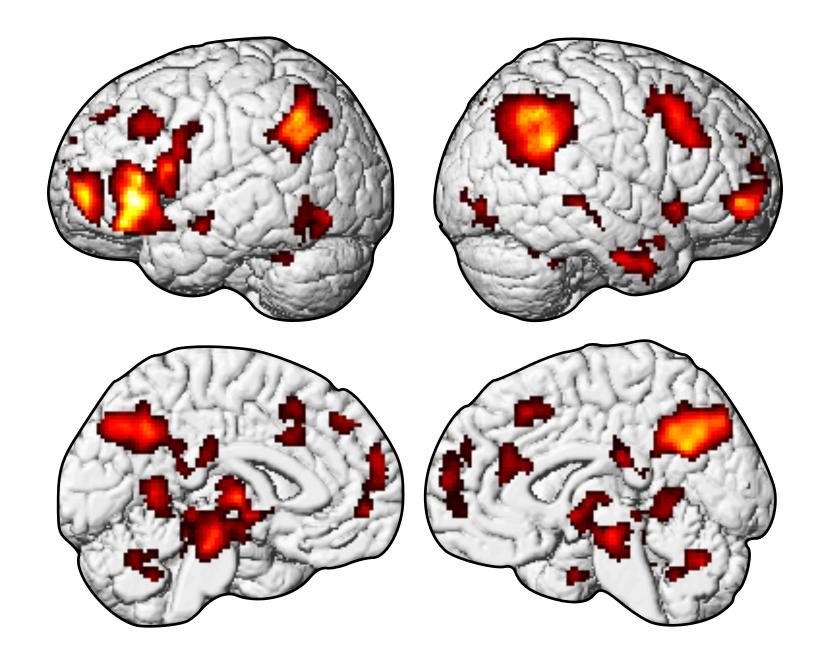


Full factorial model에서는 연구자가 F- or T- contrasts 를 정의하지 않아도, 자동으 로 가능한 모든 contrasts가 생성해 준다.

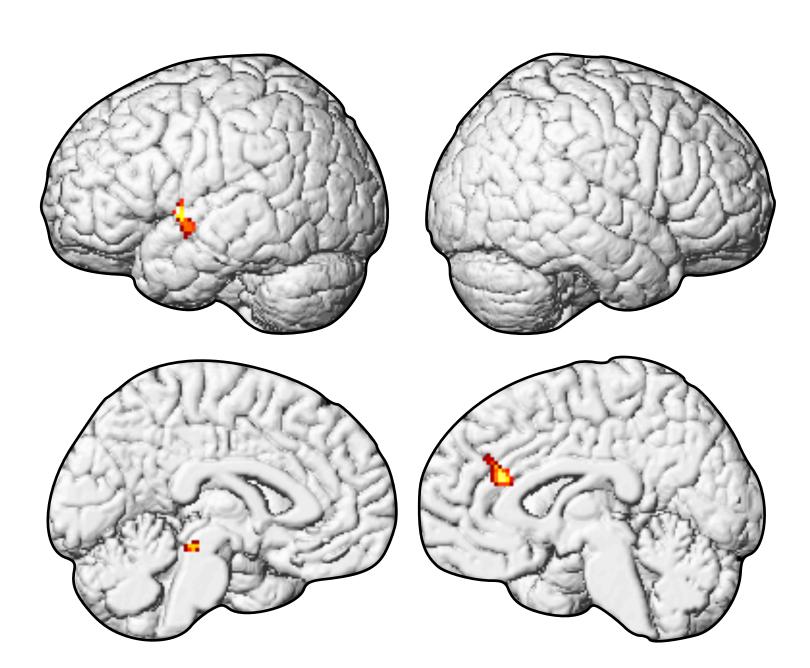
#### Full factorial: Mapping

#### Main effect of condition

Main effect of repetition



Interaction effect



CDT p<0.005 & k=50

#### 무엇을 알 수 있을까?

- Full factorial model로 데이터를 분석하게 되면, 각각의 요인에 대한 주요 효과(Main effect)와 요인과 요인 간에 상호작용 효과가 나타나는 뇌영역을 알 수 있다.
- 하지만, subject에 대한 random effects가 모델에 포함되지 않는다.

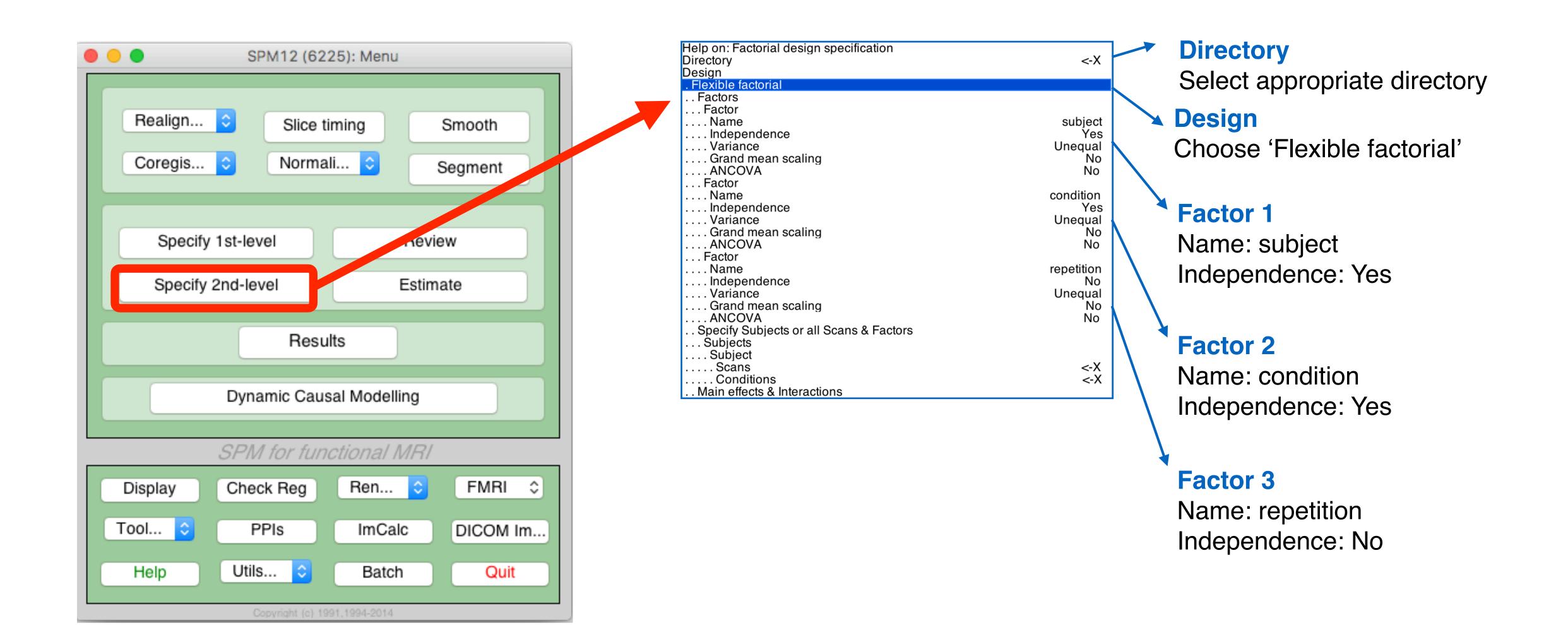
# 그룹 통계 분석 (2nd-level analysis)

- a. one sample t-test
- b. paired sample t-test
- c. full factorial
- d. flexible factorial

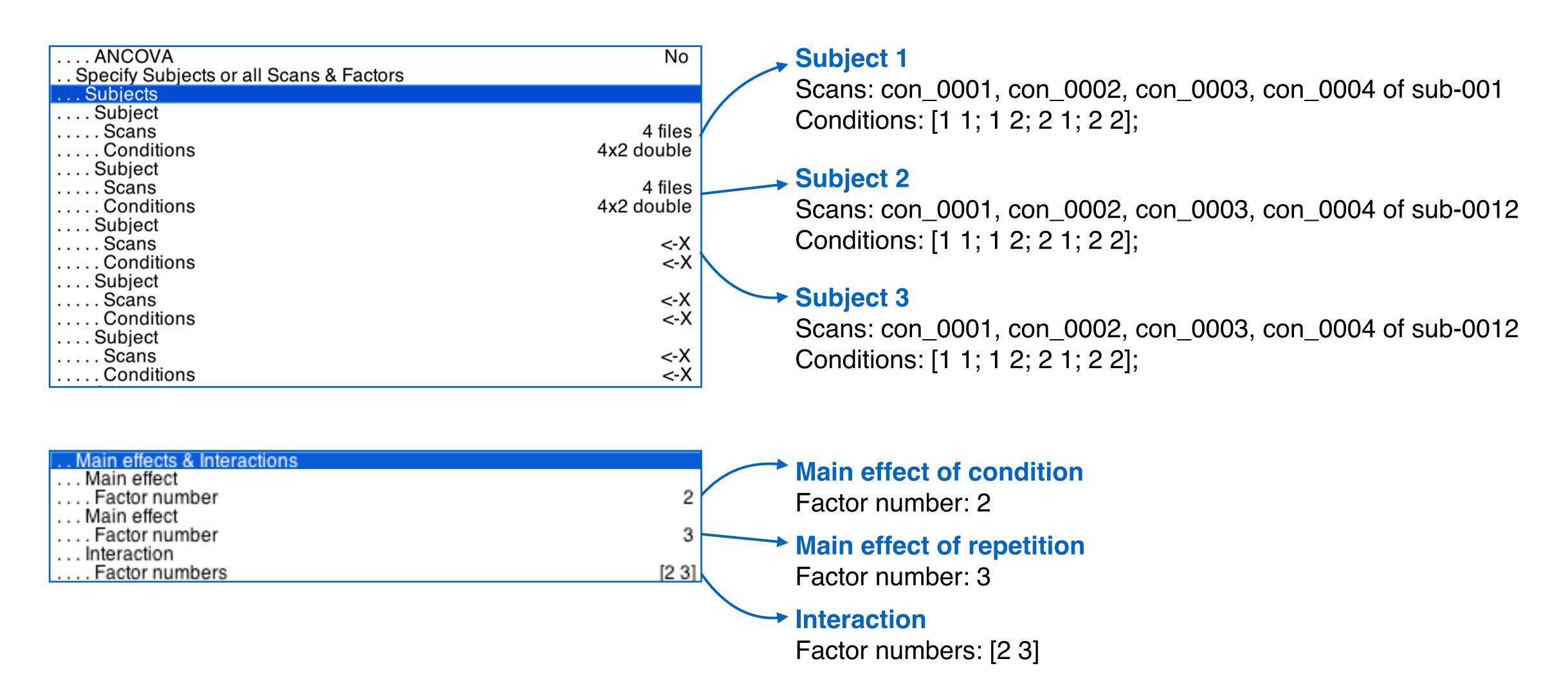
#### Flexible factorial

- This option is best used for one-way, two-way, or three-way ANOVAs but where you do not wish to test for all possible main effects and interactions.
- A typical example here would be a group-by-drug-by-task analysis where, perhaps, only (i) group-by-drug or (ii) group-by-task interactions are of interest.
   In this case it is only necessary to have two-blocks in the design matrix - one for each interaction. The three-way interaction can then be tested for using a contrast that computes the difference between (i) and (ii).
- There are two reserved words for the names of factors. There are 'subject' and 'repl' (standing for replication). If you use these factor names then SPM will automatically create replication and/or subject factors without you having to type in an extra entry in the condition vector.

## Flexible factorial: Specify model (1/2)

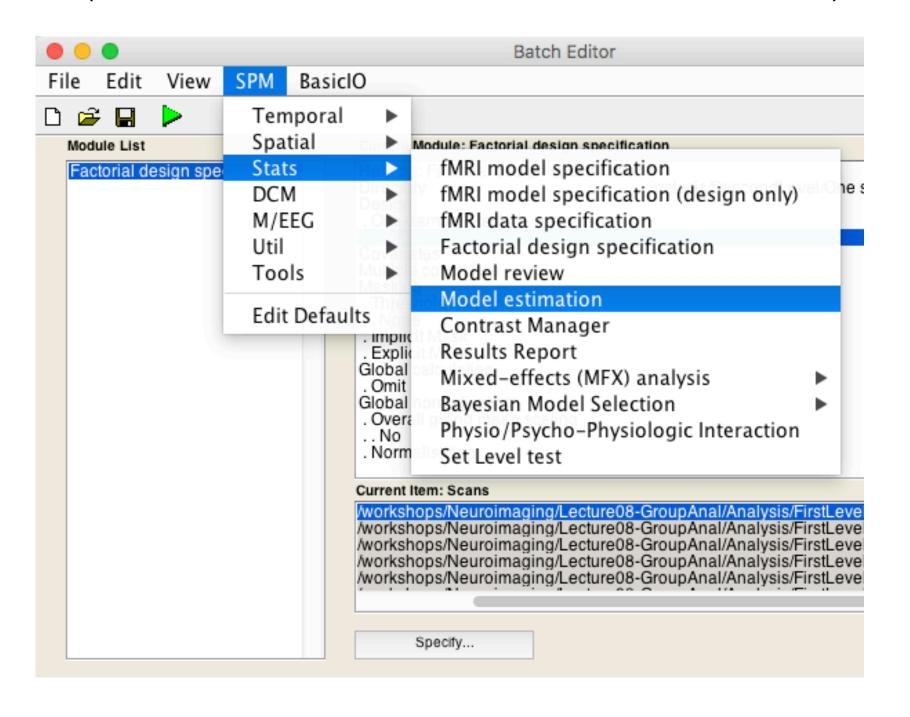


## Flexible factorial: Specify model (2/2)

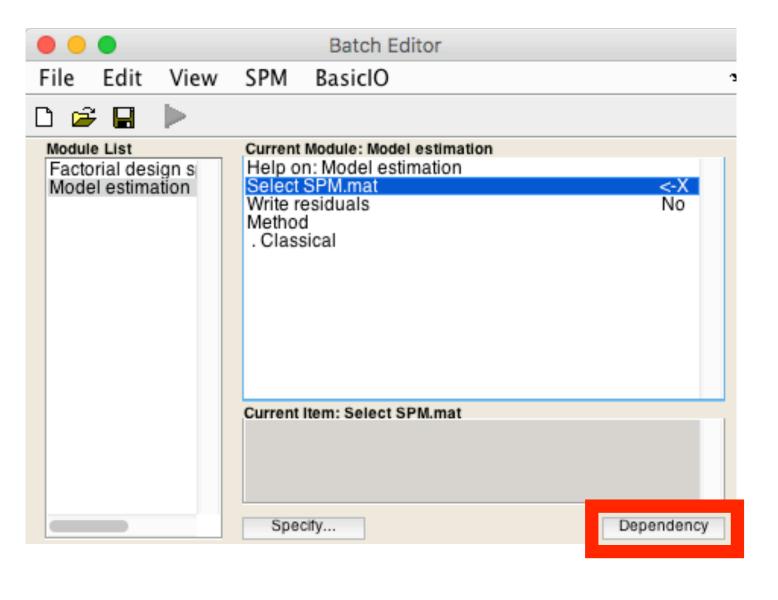


#### Flexible factorial: Estimation

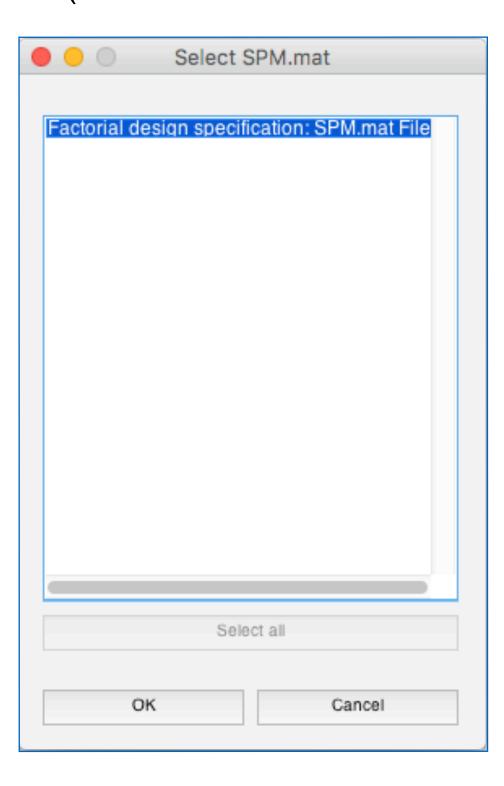
1. Add 'model estimation' in the module list (SPM menu → Stats → Model estimation)



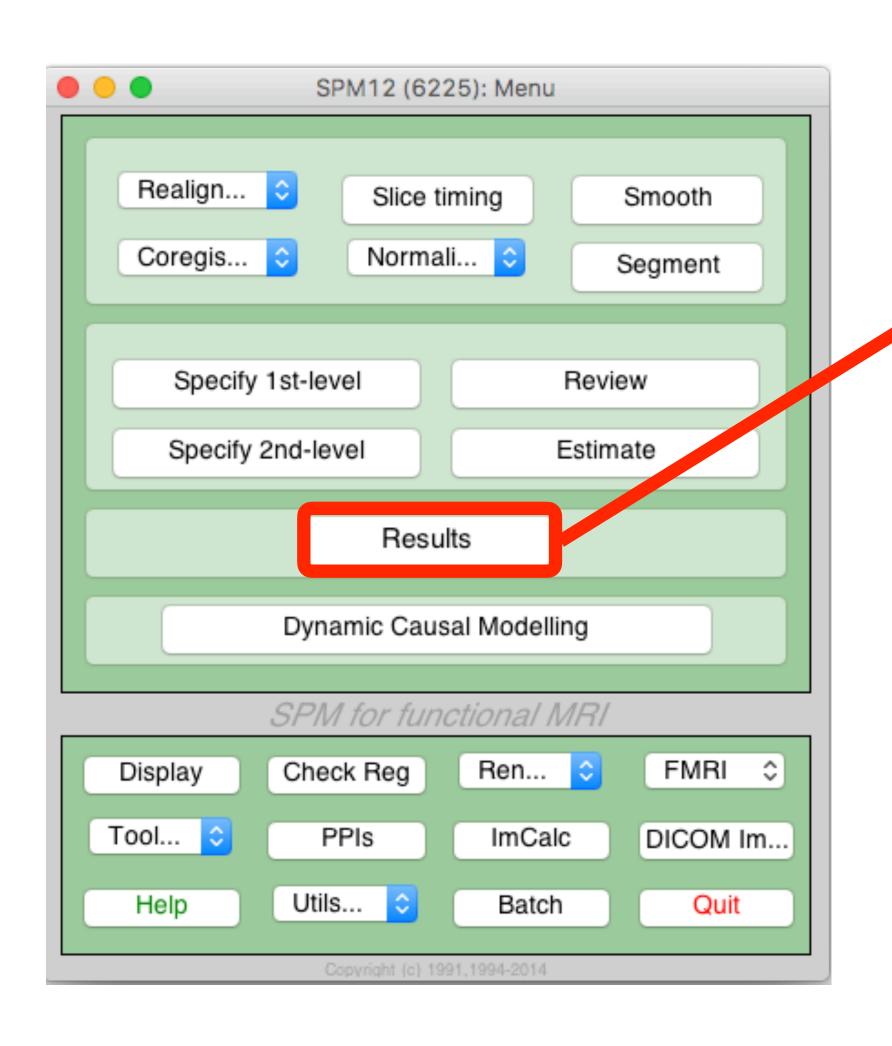
2. Select SPM.mat file (Click 'Dependency' button)



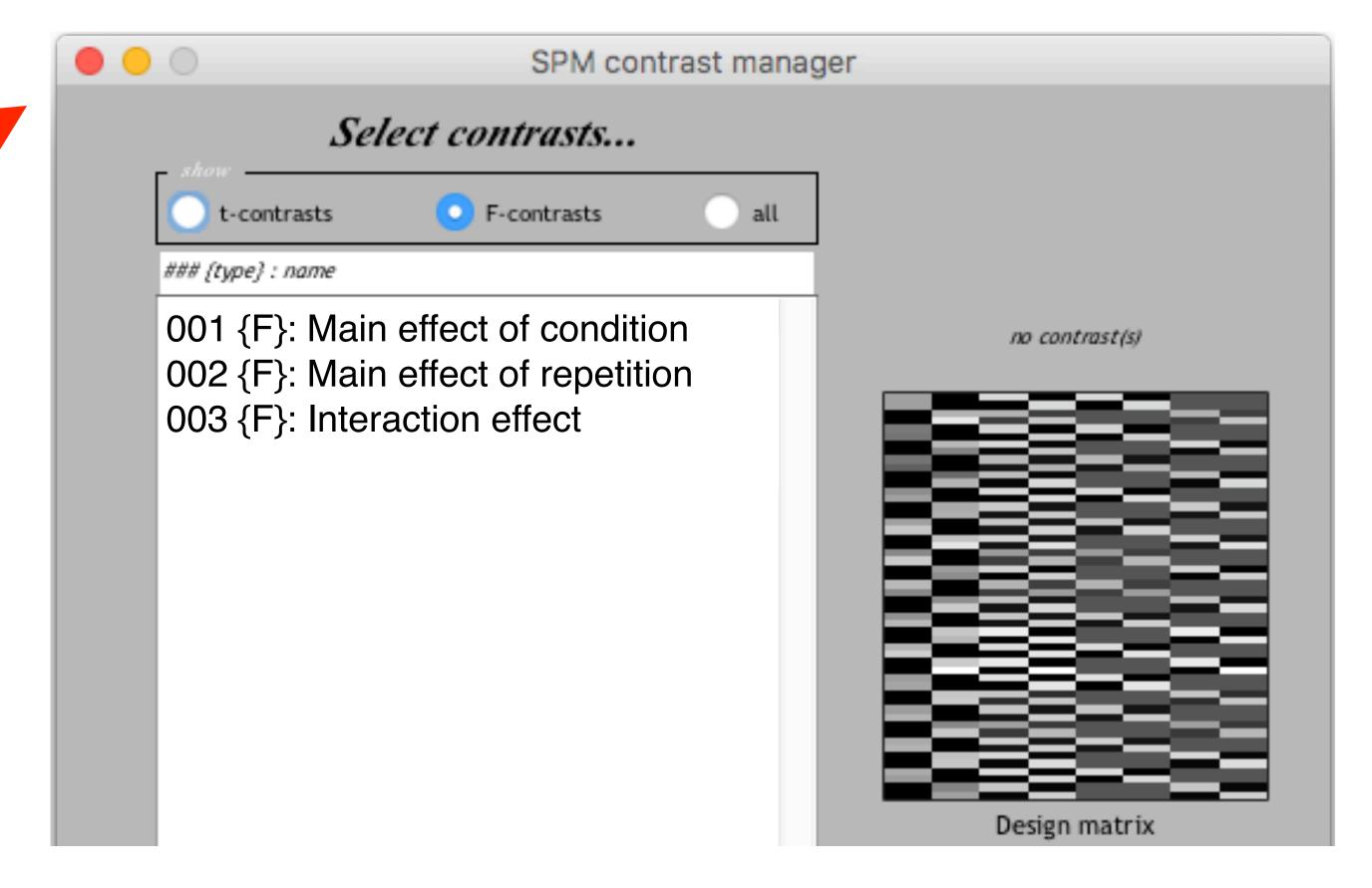
3. Select SPM.mat file (click ▶ Run after selection)



#### Flexible factorial: Contrast



Select one of existing contrasts or Define net contrast



#### More on F-contrasts

		.*	<b>©</b>		Alostrac <sup>a</sup>	18 Movel	Coucles	e Mover
Contrast name	Abstract	Concre	Money	Rebeat	Abstrac	Abstrac	Coucker	Coucke
001 {F}: Main effect of condition	1	-1	0	0	0.5	0.5	-0.5	-0.5
002 {F}: Main effect of repetition	0	0	1	-1	0.5	-0.5	0.5	-0.5
003 {F}: Interaction effect	0	0	0	0	1	-1	-1	1

#### 다양한 F-contrast에 대한 예시

Contrast weights in flexible factorial design with multiple groups of subjects

Jan Gläscher<sup>1</sup> and Darren Gitelman<sup>2</sup>

Div. of Humanities and Social Science, Caltech Dept. of Neurology and Radiology, Northwestern

March 18, 2008

contact: glascher@hss.caltech.edu

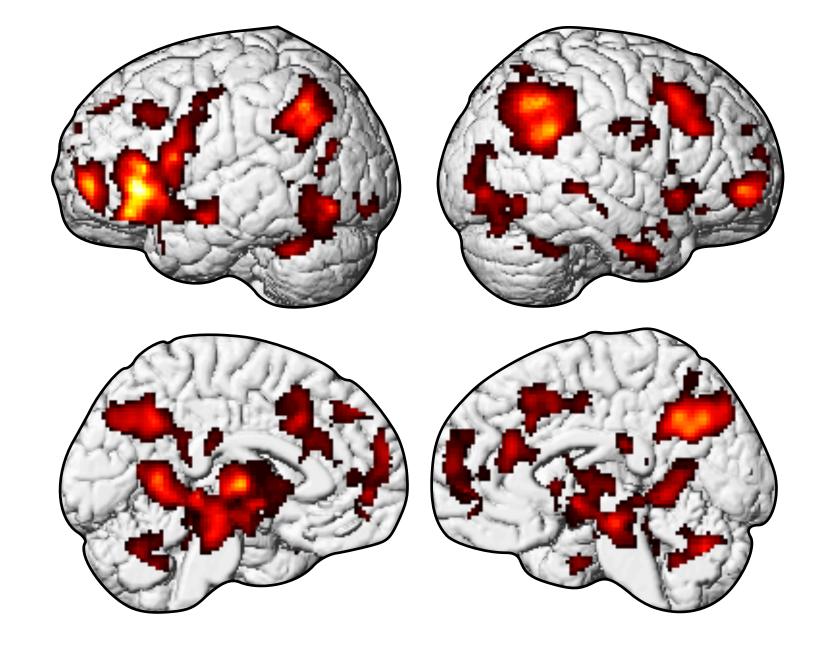
## Matlab script for flexible factorial analysis

```
>> % Directory containing Cross-language data
                                                                                                  >> % Parameter Estimation
                                                                                                  >> %------
>> proj path = '/Volumes/JetDrive/workshops/Neuroimaging/Lecture08-GroupAnal';
                                                                                                  >> SPM mat = fullfile(out dir, 'SPM.mat');
                                                                                                  >> matlabbatch{2}.spm.stats.fmri est.spmmat = {SPM mat};
>> fn xls = fullfile(proj path, 'subjlist.xlsx');
                                                                                                  >> matlabbatch{2}.spm.stats.fmri est.write residuals = 0;
>>
                                                                                                  >> matlabbatch{2}.spm.stats.fmri est.method.Classical = 1;
>> T = readtable(fn xls);
>> subjlist = T.subjname; nsubj = length(subjlist);
                                                                                                  >>
                                                                                                  >> % Create Contrasts
>>
>> % Specify Model
                                                                                                  >> matlabbatch{3}.spm.stats.con.spmmat = {SPM mat};
                                                                                                  >> matlabbatch{3}.spm.stats.con.consess{1}.fcon.name = 'Main effect of condition';
>> clear matlabbatch;
                                                                                                  >> matlabbatch{3}.spm.stats.con.consess{1}.fcon.weights = [1 -1 0 0 0.5 0.5 -0.5 -0.5];
>> out dir = fullfile(proj path, 'Analysis', 'SecondLevel', 'Flexible'); mkdir(out dir);
                                                                                                  >> matlabbatch{3}.spm.stats.con.consess{1}.fcon.sessrep = 'none';
>>
                                                                                                  >> matlabbatch{3}.spm.stats.con.consess{2}.fcon.name = 'Main effect of repetition';
>> matlabbatch{1}.spm.stats.factorial design.dir = {out dir};
                                                                                                  >> matlabbatch{3}.spm.stats.con.consess{2}.fcon.weights = [0 0 1 -1 0.5 -0.5 0.5 -0.5];
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.fac(1).name = 'subject';
                                                                                                  >> matlabbatch{3}.spm.stats.con.consess{2}.fcon.sessrep = 'none';
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.fac(1).dept = 0;
                                                                                                  >> matlabbatch{3}.spm.stats.con.consess{3}.fcon.name = 'Interaction';
>> matlabbatch{1}.spm.stats.factorial_design.des.fblock.fac(1).variance = 1;
                                                                                                  >> matlabbatch{3}.spm.stats.con.consess{3}.fcon.weights = [0 0 0 0 1 -1 -1 1];
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.fac(2).name = 'condition';
                                                                                                  >> matlabbatch{3}.spm.stats.con.consess{3}.fcon.sessrep = 'none';
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.fac(2).dept = 0;
                                                                                                  >> matlabbatch{3}.spm.stats.con.delete = 1;
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.fac(2).variance = 1;
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.fac(3).name = 'repetition';
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.fac(3).dept = 1;
                                                                                                  >> % spm jobman('interactive', matlabbatch);
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.fac(3).variance = 1;
                                                                                                  >> spm jobman('run', matlabbatch);
>>
>>
>> for c=1:nsubj,
       confiles = cell(0);
      confiles{1} = fullfile( proj path, 'Analysis', 'FirstLevel', subjlist{c}, 'con 0001.nii');
>>
      confiles{2} = fullfile( proj_path, 'Analysis', 'FirstLevel', subjlist{c}, 'con_0002.nii');
>>
      confiles{3} = fullfile( proj_path, 'Analysis', 'FirstLevel', subjlist{c}, 'con_0003.nii');
>>
      confiles{4} = fullfile( proj path, 'Analysis', 'FirstLevel', subjlist{c}, 'con 0004.nii');
>>
>>
      matlabbatch{1}.spm.stats.factorial_design.des.fblock.fsuball.fsubject(c).scans = [confiles'];
>>
      matlabbatch{1}.spm.stats.factorial_design.des.fblock.fsuball.fsubject(c).conds = [1 1; 1 2; 2 1; 2 2];
>>
>>
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.maininters{1}.fmain.fnum = 2;
>> matlabbatch{1}.spm.stats.factorial design.des.fblock.maininters{2}.fmain.fnum = 3;
>> matlabbatch{1}.spm.stats.factorial_design.des.fblock.maininters{3}.inter.fnums = [2 3];
```

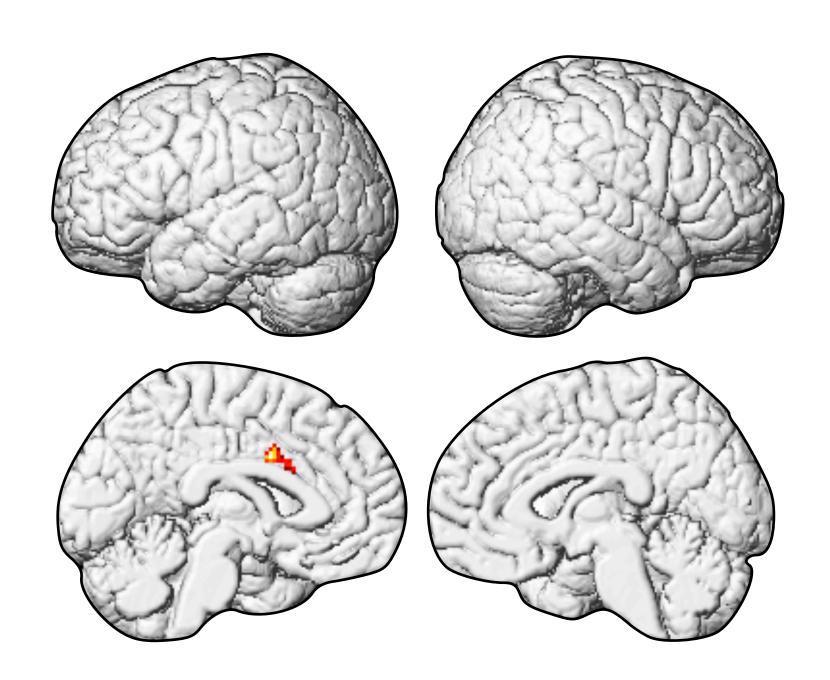
#### Flexible factorial: Mapping

#### Main effect of condition

Main effect of repetition



Interaction effect



CDT p<0.005 & k=50