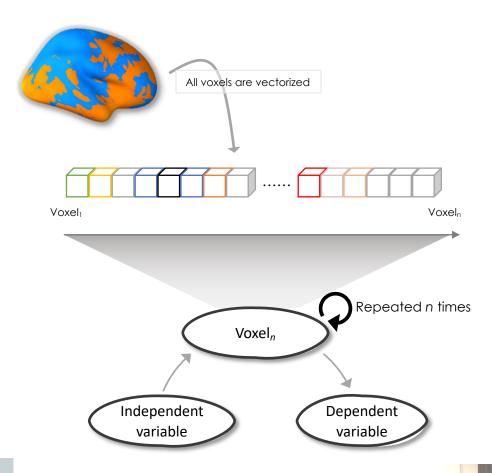
#### mediation\_dream

(https://github.com/cocoanlab/cocoanCORE)

Motivation 1:

: When performing multilevel mediation analysis for whole brain, It takes a lot of time to get results because only one CPU core is working for the analysis

e.g., if you have 200,000 voxels, the analysis will be performed 200,000 times sequentially



# It usually takes two weeks to finish this analysis



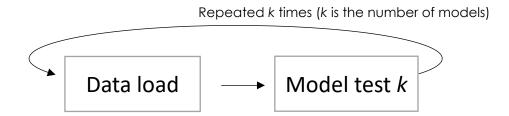
#### mediation\_dream

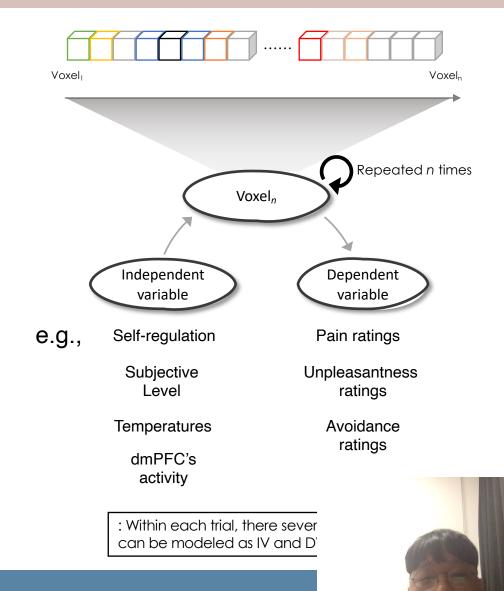
(https://github.com/cocoanlab/cocoanCORE)

Motivation 2:

: If we want to test several models, it is also hugely influenced by the number of the models.

: Whenever the models are tested, same fMRI data should be loaded repeatedly



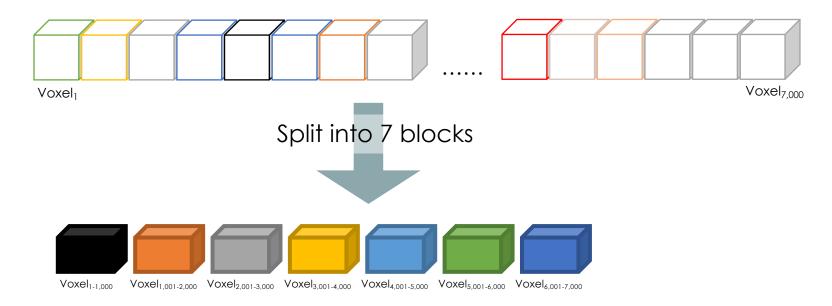




- mediation\_dream (<a href="https://github.com/cocoanlab/cocoanCORE">https://github.com/cocoanlab/cocoanCORE</a>)
  - To overcome these issues (\*WE WANT TO GET ANALYSIS RESUTLS FASTER\*), this function was created
    - You can reduce a processing time from 2 weeks to 3 or 4 days
    - It is intended to performing parallel processing of whole-brain multilevel mediation analysis
  - CanlabCORE, cocoanCORE and mediationToolbox must be added to your path
  - Recommendations
    - It is for the multilevel whole—brain mediation analysis especially,
      - with bootstrap option
      - voxels > 1,000 and models > 2
    - A high-performance computer with several CPU cores and RAM storage is needed



- The algorithm of mediation\_dream
  - If you want to perform seven simultaneous the whole-brain mediation analysis
    - 1. All voxels in brain are divided into **7 blocks**



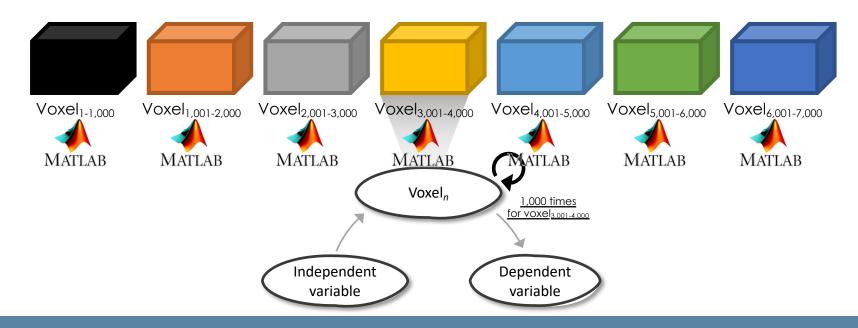
- The algorithm of mediation\_dream
  - If you want to perform seven simultaneous the whole-brain mediation analysis
    - 1. All voxels in brain are divided into 7 blocks
    - 2. Put all models you want to test in the function

	IV	DV			
Model 1	Self-regulation	Pain ratings			Repeated <i>k</i> times ( <i>k</i> is the number of models)
Model 2	Subjective Level	Unpleasantness ratings		]	
Model 3	Temperatures	Avoidance ratings	Data load		Model test k
Model 4	dmPFC's activity	Pain ratings			



#### The algorithm of mediation\_dream

- If you want to perform seven simultaneous the whole-brain mediation analysis
  - 1. All voxels in brain are divided into **7 blocks**
  - 2. Put all models you want to test in the function
  - 3. Run each script on each CPU core in HPC

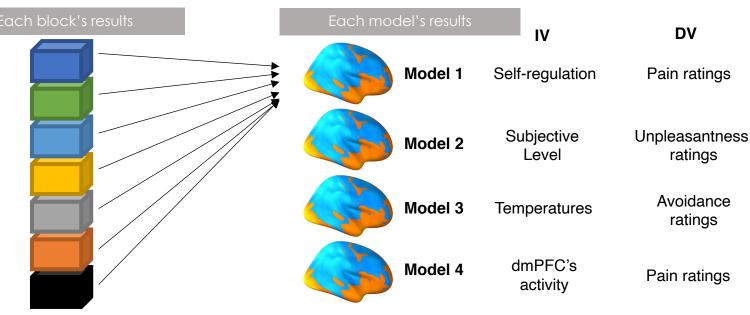




#### The algorithm of mediation\_dream

- If you want to perform seven simultaneous the whole-brain mediation analysis
  - 1. All voxels in brain are divided into 7 blocks
  - 2. Put all models you want to test in the function
  - 3. Run each script on each CPU core in HPC
  - 4. Summarize the results

For each model, the results are summarized as path a, path b, and path A\*b

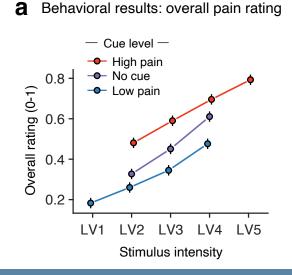




- Be careful
  - This function uses a global variable
  - Making a brain mask for own study is important
    - Even if each brain was spatially normalized, some voxels can have zero values (especially, in the brainstem)
    - If this situation happens, it can generate an error regarding no variance



- STEP1: Make variable structure
- STEP2: Build models
- STEP3: Divide fMRI data and scripts for parallel processing (mediation\_dream\_wani.m)
- STEP4: Run each script in MATLAB with each CPU core (1CPU core = 1 script)
- STEP5: Summarize each result (mediation\_dream\_combined\_wani.m)



Whole-brain activity during heat stimulation

X Cue Y Overall Rating

Covariate: stimulus intensity

Whole-brain mediation for the cue effects

Whole-brain activity during heat stimulation

X Stimulus overall Rating

Covariate: Cue level

Whole-brain mediation for the stimulus intensity

(Gim et al., in prep)



STEP1: Make variable structure

(see cocoanCORE/Statistics/mediation\_dream/mediation\_dream\_suhwan\_example\_code\_3.m)

#### The structure of 'med\_vars'

```
med_vars =

struct with fields:

    M: {1x58 cell}
    X1: {1x58 cell}
    X2: {1x58 cell}
    Y: {1x58 cell}
    model_name: '/sas1/cocoanlab/data/SEMIC/analysis/imaging/first_level/model03a_
```

med\_vars.X1 = stimulus intensity med\_vars.X2 = cue level med\_vars.Y = overall pain rating med\_vars.M = the address of heat-stimulation induced brain activity (fMRI data\_

#### The address of fMRI data should be included as cell type

```
med_vars.M{1}
ans =

66×1 cell array

{'/sas1/cocoanlab/data/SEMIC/analysis/imaging/first_level/model03a_SPM_SINGLE_TRIAL_PAIN/sub-semic001/beta_0001.nii'}
 {'/sas1/cocoanlab/data/SEMIC/analysis/imaging/first_level/model03a_SPM_SINGLE_TRIAL_PAIN/sub-semic001/beta_0002.nii'}
 {'/sas1/cocoanlab/data/SEMIC/analysis/imaging/first_level/model03a_SPM_SINGLE_TRIAL_PAIN/sub-semic001/beta_0003.nii'}
```



STEP2: Build models

You build model to test in models.name{k} and models.fns{k}

```
models.name{1} = 'model01_X:_Stim_Y:_angle_M:_brain_Cov:_cue';
models.name{2} = 'model02_X:_Cue_Y:_angle_M:_brain_Cov:_Stim';|

models.fns{1} = ['mediation(med_vars.X1, med_vars.Y, M,''cov'', med_vars.X2, ''boot'', ''bootsamples'', 10000)'];
models.fns{2} = ['mediation(med_vars.X2, med_vars.Y, M,''cov'', med_vars.X1, ''boot'', ''bootsamples'', 10000)'];

>> disp(models.fns{1})
mediation(med_vars.X1, med_vars.Y, M,'covs', med_vars.X2, 'boot', 'bootsamples', 10000)
>> disp(models.fns{2})
mediation(med_vars.X2, med_vars.Y, M,'covs', med_vars.X1, 'boot', 'bootsamples', 10000)
```

k = the number of model

The mediation function you want to perform



STEP3: Divide fMRI data and scripts for parallel processing (mediation\_dream\_wani.m)
 Run mediation\_dream\_wani.m (or \_suhwan.m)

The number of job you want to divide

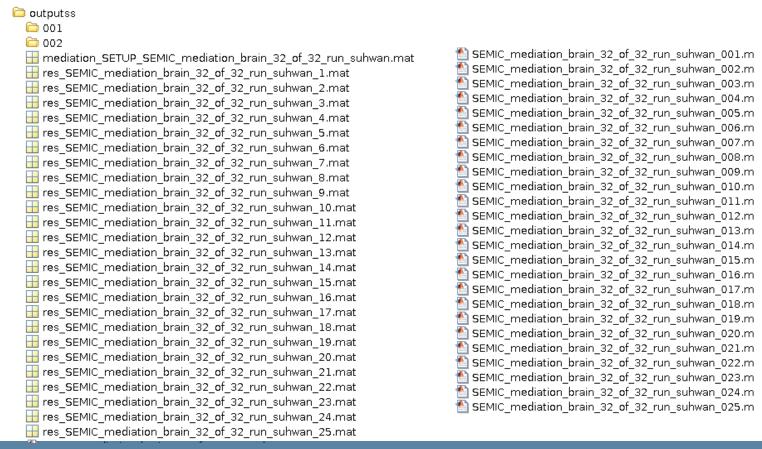
```
For specific
                           jobn = 26;
environment
                          wh_loc = 'HPC';
                          % it should change directory or filename
                          outputdir = fullfile('/sas1/cocoanlab/data','SEMIC','191203_2mm_whole_brain_onlySELF','outputs');
      med_vars.imgs
                          if ~exist(outputdir, 'dir'), mkdir(outputdir); end
                          med_vars.imgs = med_vars.M;
                          %models.name{1} = sprintf('Model01_STIM_phase_%02d_of_08_whole_brain',sec_i);
                          %models.name{2} = sprintf('Model02 CUE phase %02d of 08 whole brain', sec i);
                          for i = 1:length(models.fns)
                              models.savepaths{i} = [1,2,3,4,5]; % including all path
                          end
                          code_filename = fullfile(outputdir, sprintf('SEMIC_mediation_brain_%02d_of_32_run_suhwan.m',sec_i));
                          study_scriptdir = fullfile('/sas1/cocoanlab/data/SEMIC','scripts');
                          %% make mediation distributed scripts
                          mediation_dream_suhwan(med_vars, models, jobn, mask, code_filename, study_scriptdir,'wh_loc',wh_loc)
```

This function is for my specific environment
You can use \_wani or modify it for your environment



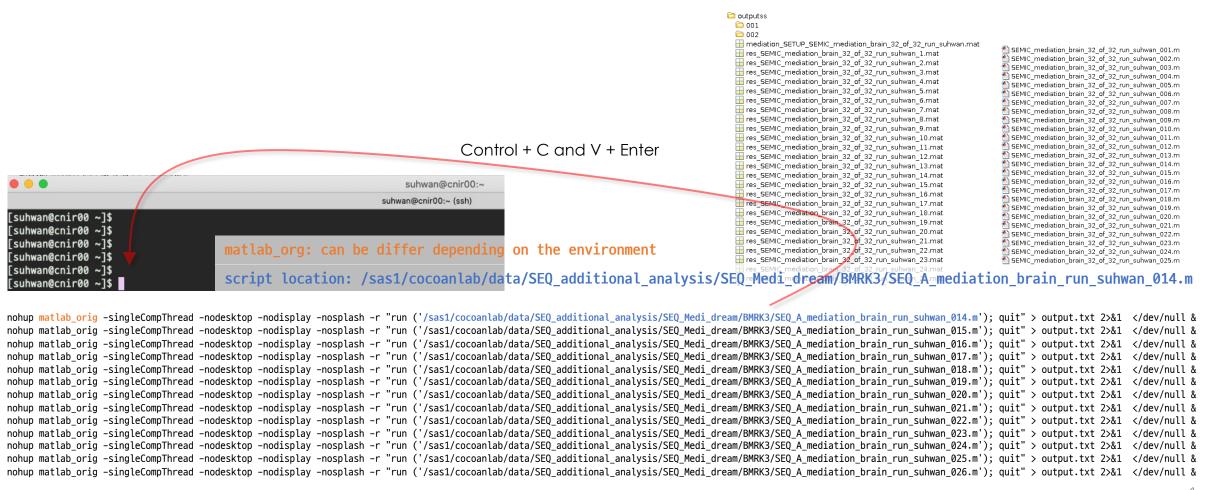
STEP4: Run each script in MATLAB with each CPU core (1CPU core = 1 script)

#### These data (.mat) and scripts (.m) was generated





STEP4: Run each script in MATLAB with each CPU core (1CPU core = 1 script)





STEP5: Summarize each result (mediation\_dream\_combined\_wani.m)

```
%modeldir = fullfile('/sas1/cocoanlab/data/SEMIC/','191023_2mm_whole_brain_onlySELF','outputss');
modeldir = fullfile('/sas1/cocoanlab/data/SEMIC/','191203_2mm_whole_brain_onlySELF','outputs');
sec_i = 32;
SETUP_dir = fullfile(modeldir,sprintf('mediation_SETUP_SEMIC_mediation_brain_%02d_of_32_run_suhwan.mat',sec_i));
mediation_dream_combine_results_suhwan(modeldir, SETUP_dir);

This function is for my specific environment
You can use _wani or modify it for your environment
```



STEP5: Summarize each results (mediation\_dream\_combined\_wani.m)

```
%modeldir = fullfile('/sas1/cocoanlab/data/SEMIC/','191023 2mm whole brain onlySELF','outputss');
modeldir = fullfile('/sas1/cocoanlab/data/SEMIC/','191203_2mm_whole_brain_onlySELF','outputs');
sec i = 32;
SETUP_dir = fullfile(modeldir,sprintf('mediation_SETUP_SEMIC_mediation_brain_%02d_of_32_run_suhwan.mat',sec_i));
mediation_dream_combine_results_suhwan(modeldir, SETUP_dir);
                                                                  You can get combined results for each model
                   001
                                                                                   002
                       📄 firstlevel M-Ybetas.nii
                                                                                      in firstlevel M-Ybetas.nii
                        firstlevel X-Mbetas.nii
                                                                                      irstlevel X-Mbetas.nii
                        firstlevel X-M-Ybetas.nii
                                                                                      📄 firstlevel X-M-Ybetas.nii
                      📄 firstlevel_X-Y_directbetas.nii
                                                                                      firstlevel X-Y directbetas.nii
                        firstlevel X-Y totalbetas.nii
                                                                                      📄 firstlevel X-Y totalbetas.nii
                        M-Y effect.nii
                                                                                      M-Y effect.nii
                        M-Y pvals.nii
                                                                                      📄 M-Y pvals.nii
                                                                                                                                a Whole-brain mediation for the cue effects
                        M-Y ste.nii
                                                                                      📄 M-Y ste.nii
                       🗋 X-M effect.nii
                                                                                      📄 X-M effect.nii
                                                                                                                                             Whole-brain activity
                        X-M pvals.nii
                                                                                      📄 X-M_pvals.nii
                                                                                                                                             during heat stimulation
                                              Mediation effects
                        X-M ste.nii
                                                                                      📄 X-M ste.nii
                       X-M-Y effect.nii
                                                                                      📄 X-M-Y effect.nii
                        X-M-Y pvals.nii
                                                                                      📄 X-M-Y pvals.nii
                                                                                                                                                           Overall
                        X-M-Y ste.nii
                                                                                      📄 X-M-Y ste.nii
                        X-Y direct effect.nii
                                                                                      X-Y direct effect.nii
                                                                                                                                 Covariate: stimulus intensity
                        X-Y direct pvals.nii
                                                                                      X-Y direct pvals.nii
                        X-Y direct ste.nii
                                                                                      X-Y direct ste.nii
                       📄 X-Y total effect.nii
                                                                                      X-Y total effect.nii
                      X-Y total pvals.nii
                                                                                      📄 X-Y total pvals.nii
                      X-Y total ste.nii
                                                                                      📄 X-Y total ste.nii
```



# Mediation\_Dream: Summary of example

- STEP1: Make variable structure
- STEP2: Build models
- STEP3: Divide fMRI data and scripts for parallel processing (mediation\_dream\_wani.m)
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