



## Practical Intensity Based Meta-Analysis

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# Practical intensity-based meta-analysis

Coordinate-based meta-analysis



Image-based meta-analysis



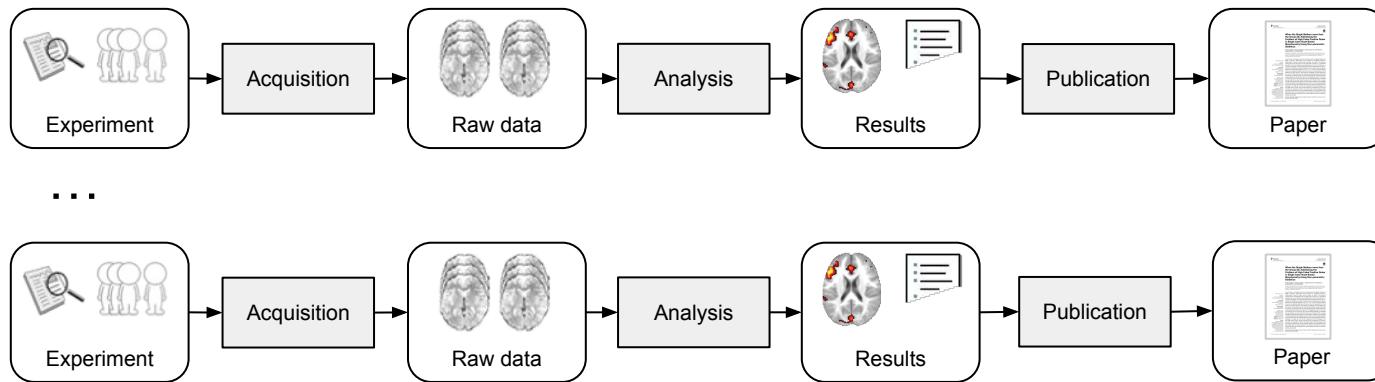
Camille Maumet

Presented by Thomas Nichols

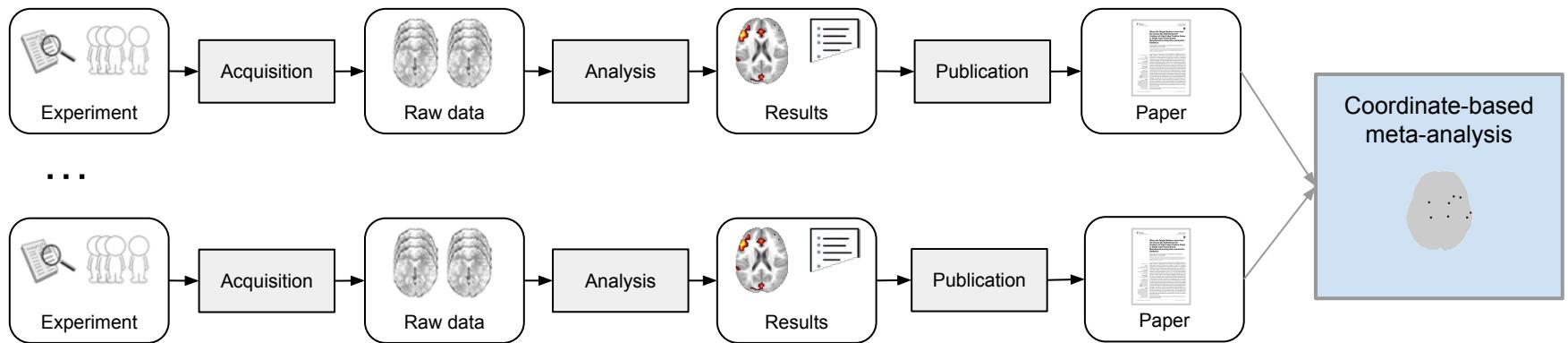
OHBM Neuroimaging Meta-Analysis Educational course  
June 17<sup>th</sup> 2018

# Coordinate-Based & Image-Based Meta-Analyses

# Neuroimaging meta-analyses



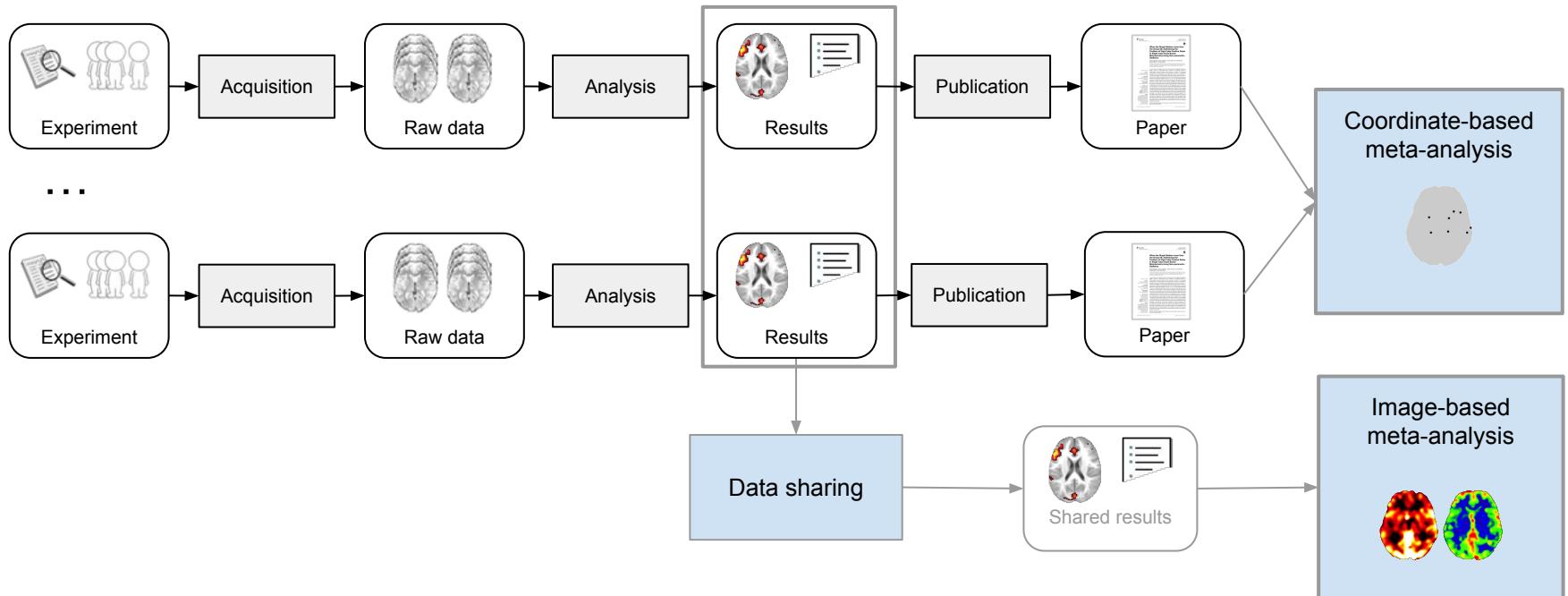
# Neuroimaging meta-analyses



## Coordinate-based meta-analysis



# Neuroimaging meta-analyses



Coordinate-based meta-analysis



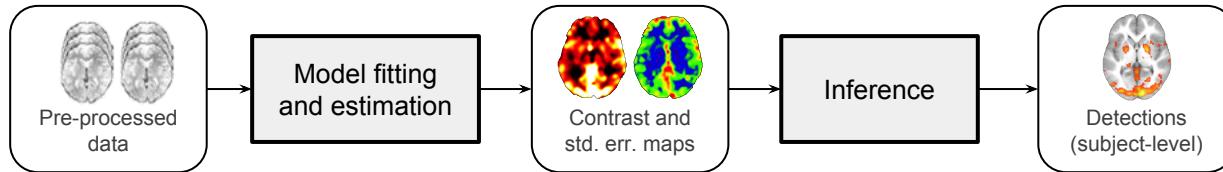
Image-based meta-analysis



How to perform an image-based  
meta-analysis?

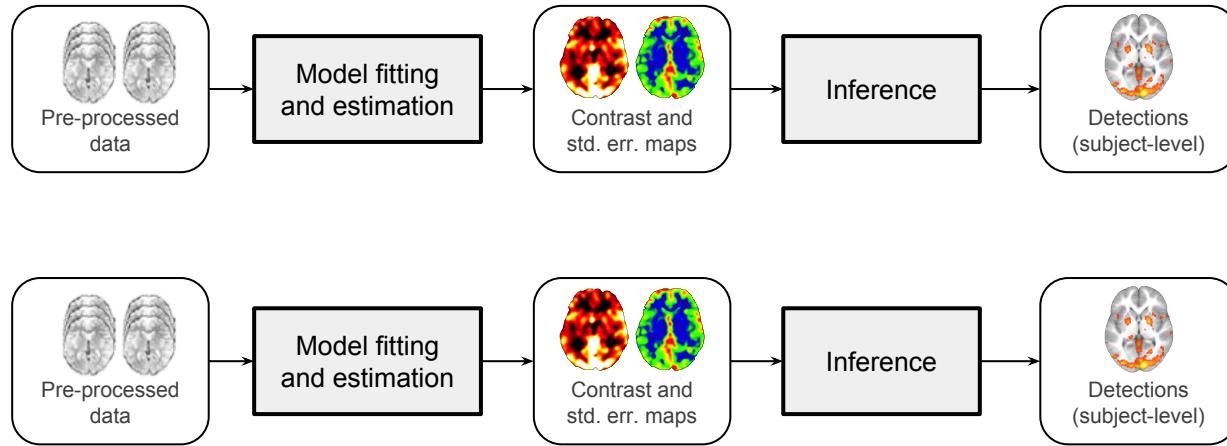
# Image-based meta-analysis

Subject 1

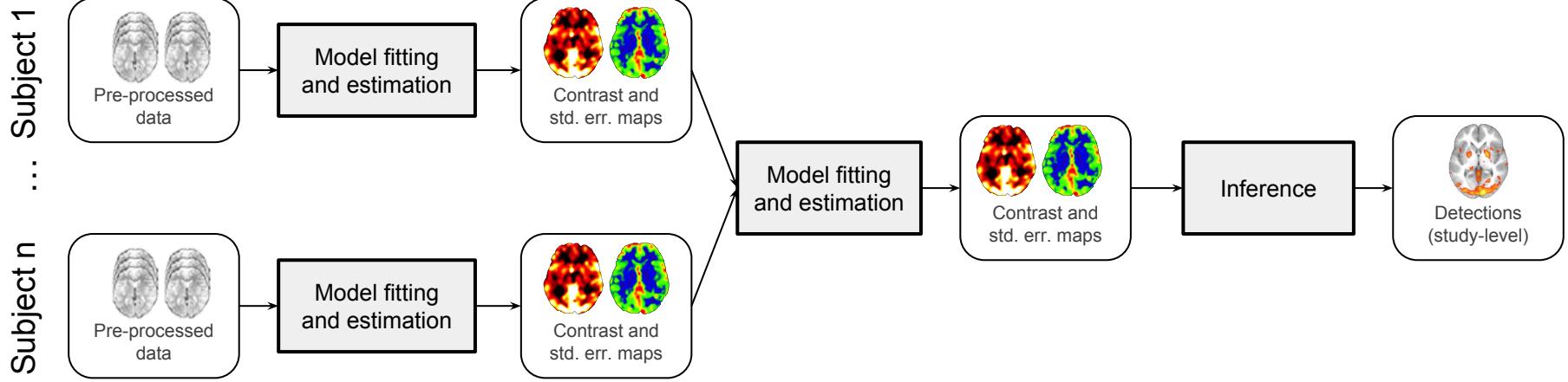


# Image-based meta-analysis

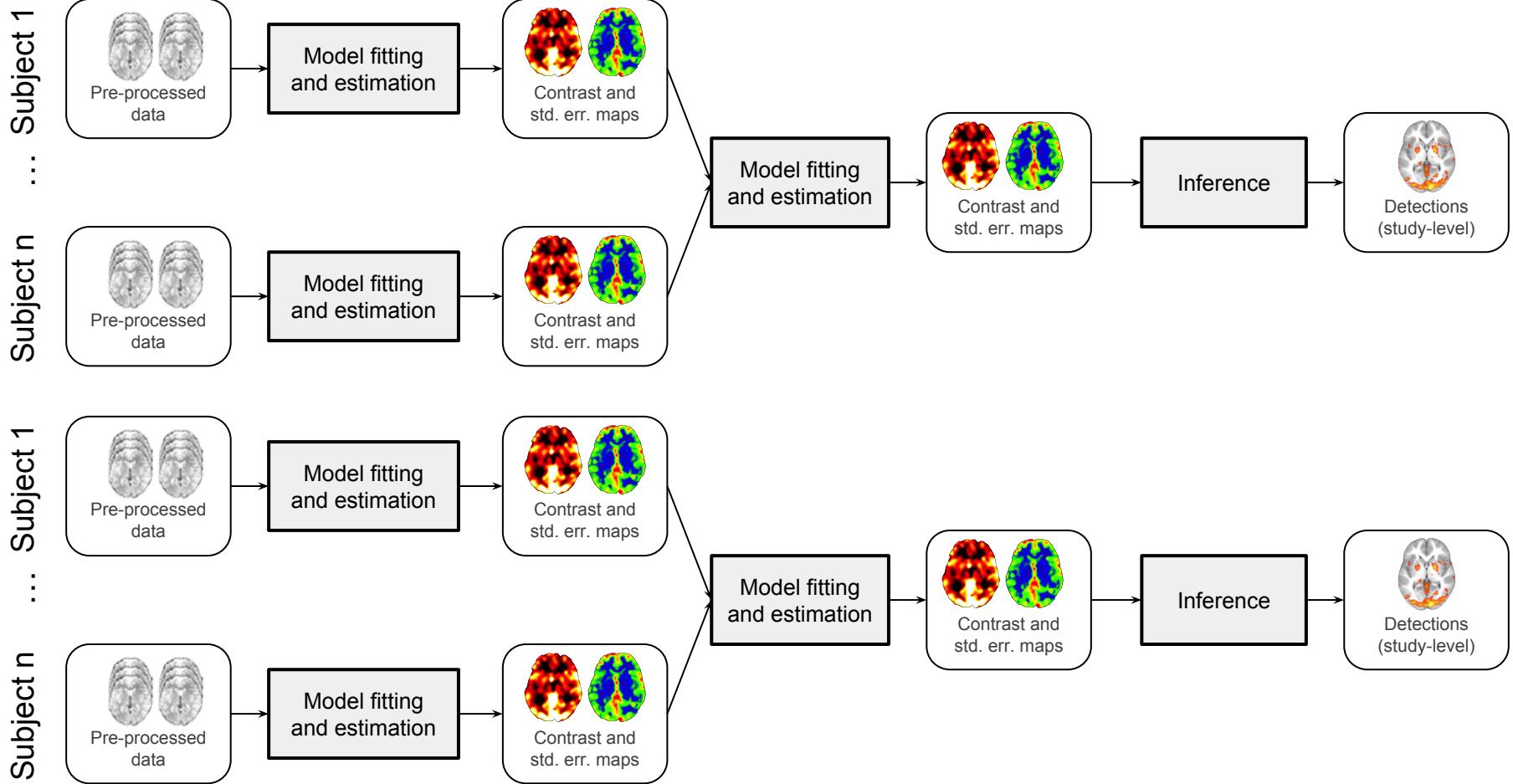
Subject 1  
...  
Subject n



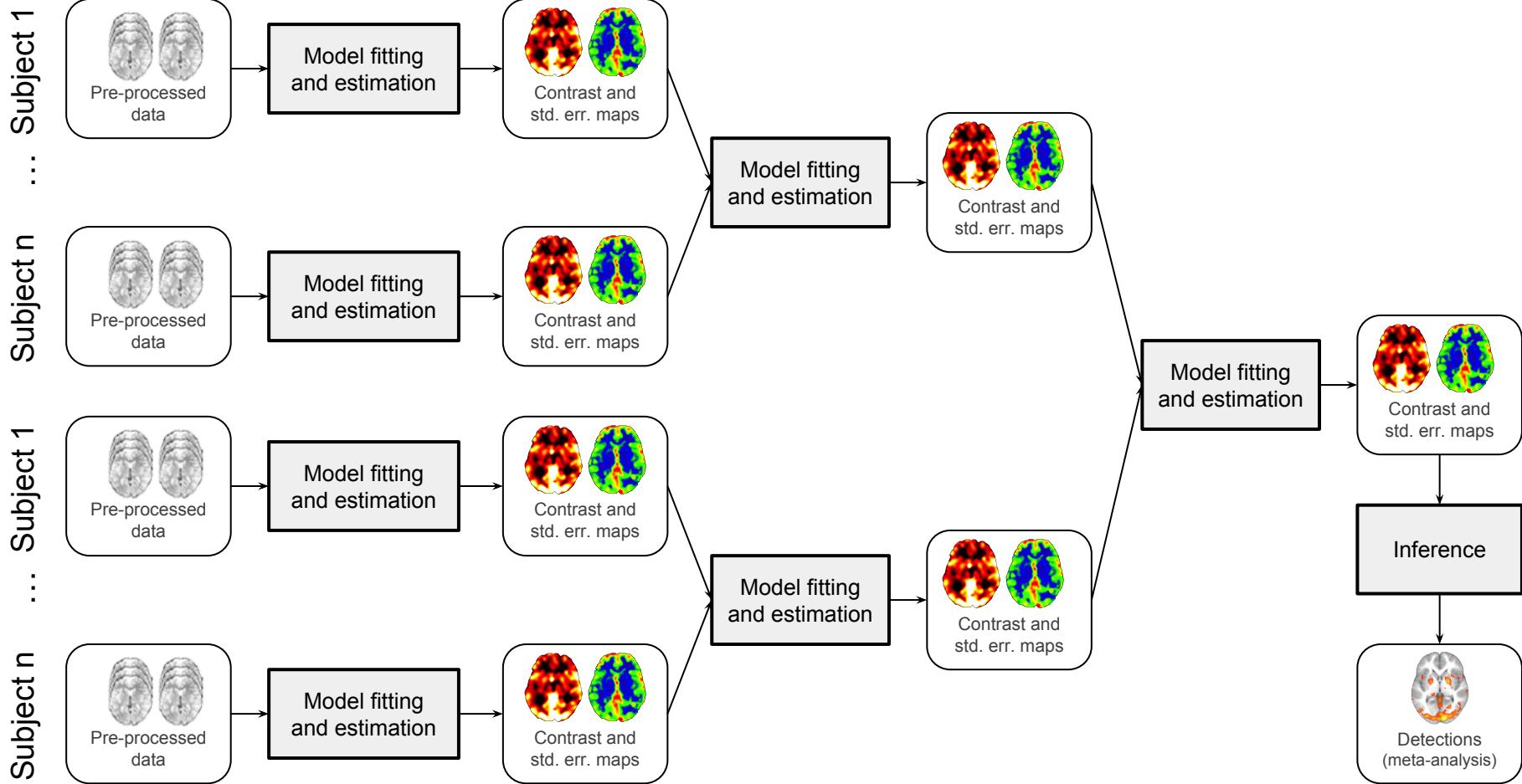
# Image-based meta-analysis



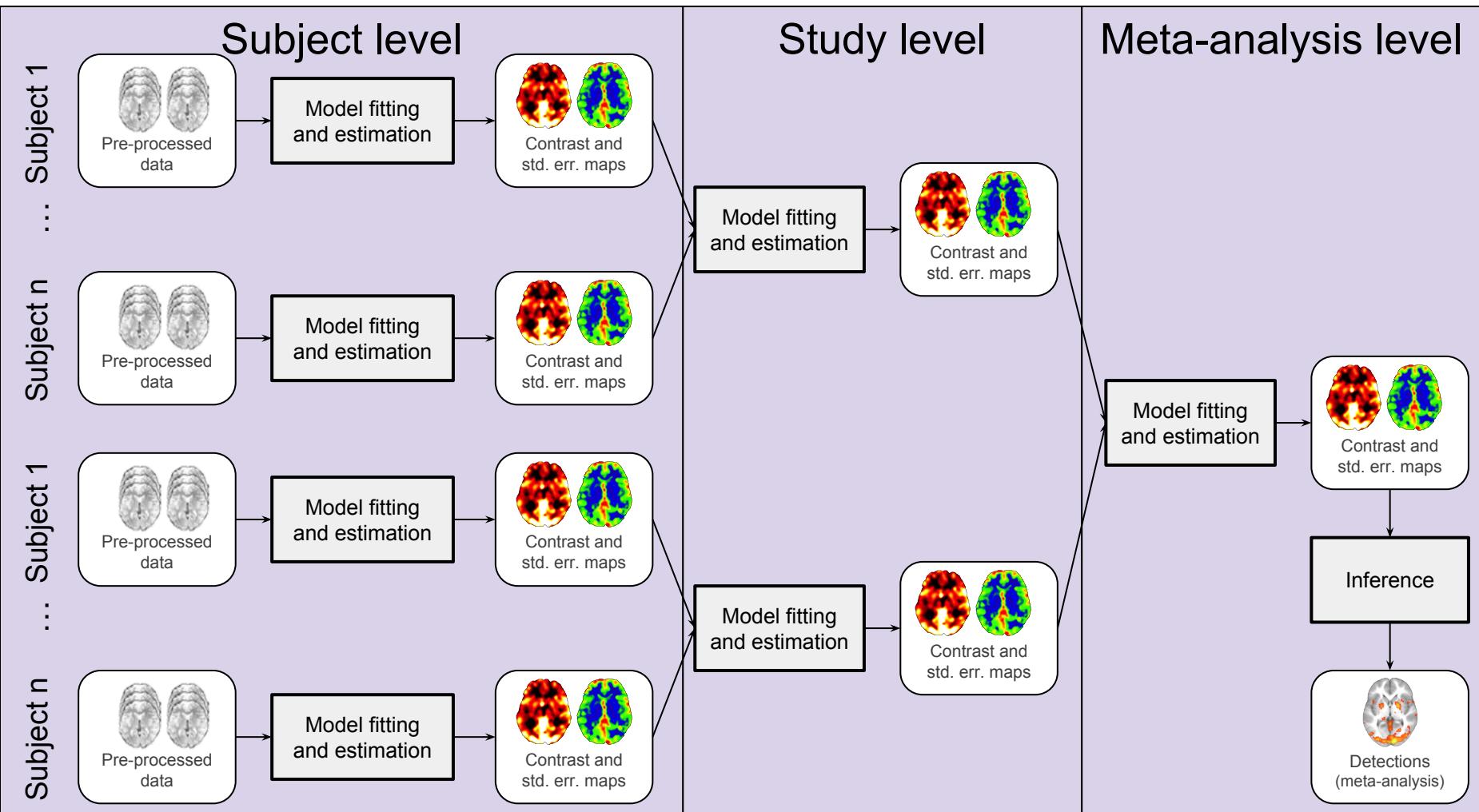
# Image-based meta-analysis



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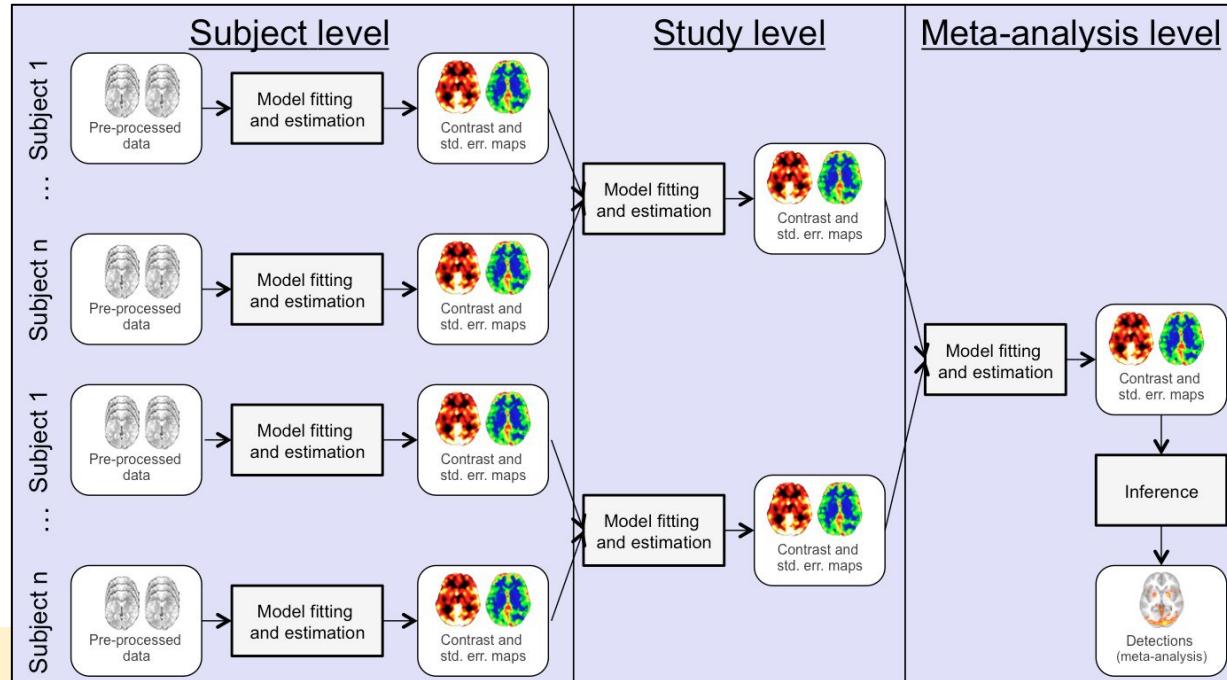
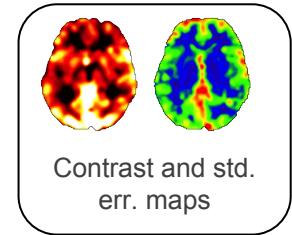
# Image-based meta-analysis

- Gold standard:

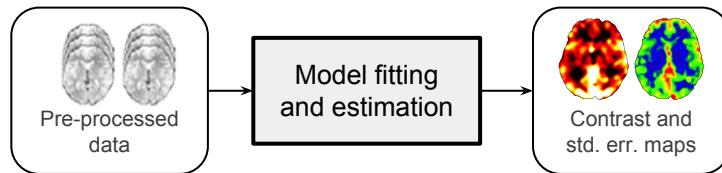
## Third-level Mixed-Effects GLM

- Requirements

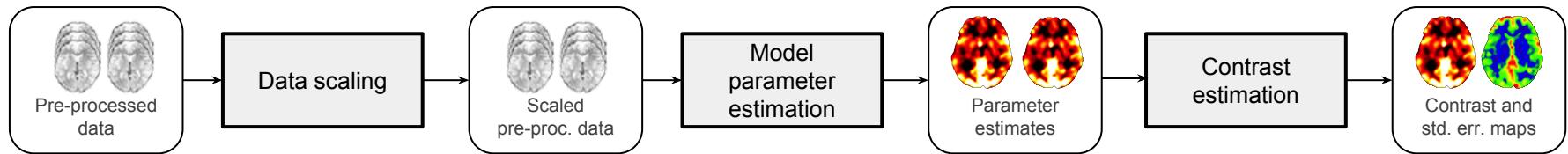
- study-level **Contrast estimates and Standard error maps.**
- **Same units**



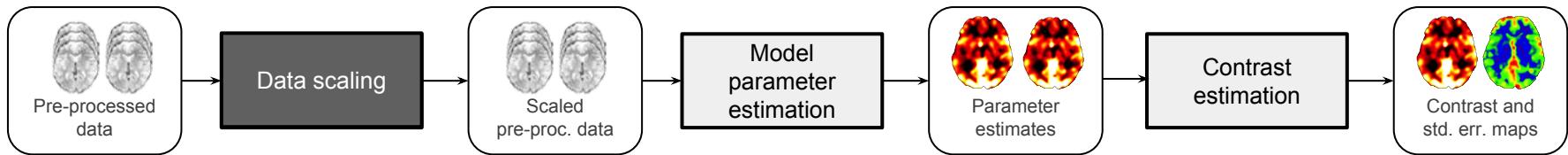
# Units of contrast estimates



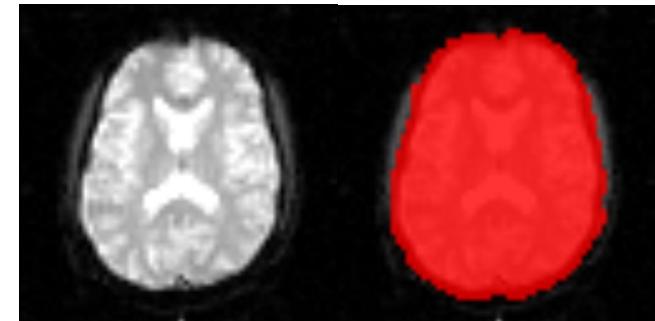
# Units of contrast estimates



# Units of contrast estimates

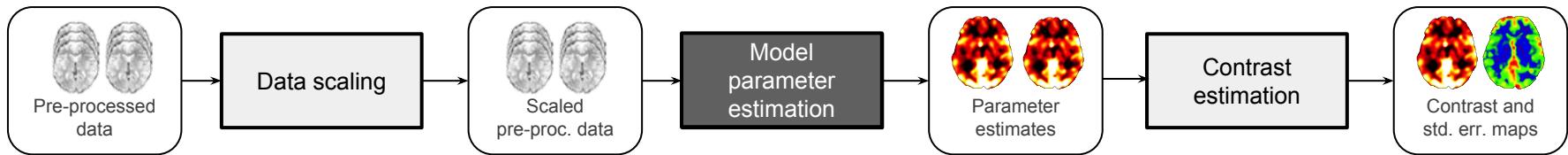


$$\text{scaled\_data} = \frac{\text{data} * \text{target}}{\text{est\_mean}}$$



Units depend on **mean estimation** and **scaling target**.

# Units of contrast estimates

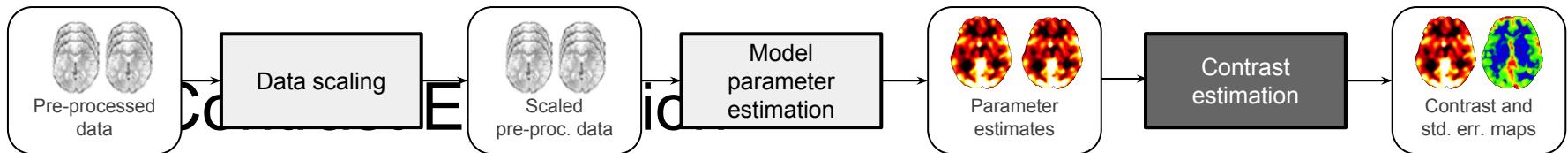


$$Y = \beta + \varepsilon$$

A large matrix diagram illustrating the linear model. The left side shows a vertical stack of 10 horizontal rows, alternating between dark gray and light gray. The right side shows a large, light gray rectangular area representing the error term  $\varepsilon$ .

Units depend on **scaling** of explanatory variables

# Units of contrast estimates



- Linear combination of parameter estimates
- Final statistics invariant to scale
  - e.g. [ 1 1 1 1 ] gives same T's & P's as [  $\frac{1}{4} \frac{1}{4} \frac{1}{4} \frac{1}{4}$  ]

Units depend on **contrast vector**

- Rule for contrasts to preserve units
  - Positive elements sum to 1
  - Negative elements sum to -1

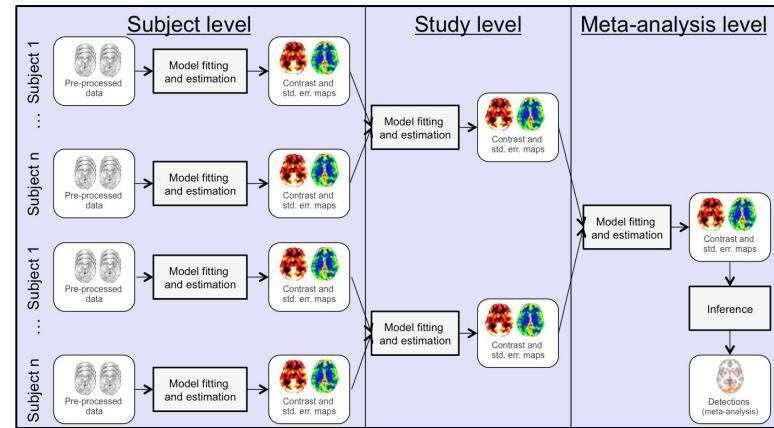
# Units of contrast estimates

- Gold standard:

## Third-level Mixed-Effects GLM

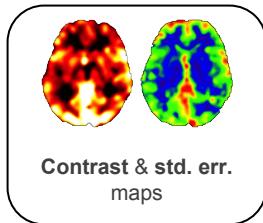
- But...

- Units will depend on:
  - The scaling of the data (subject-level)
  - The scaling of the predictor(s) (subject- and study-level)
  - The scaling of the contrast (subject- and study-level).
- Contrast estimates and standard error maps are rarely shared...



# Which images for IBMA?

**SPM**



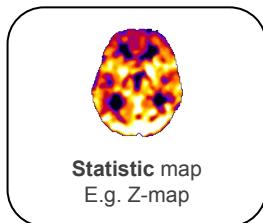
con\_0001.nii  
[SPM.mat]



cope1.nii  
varcope1.nii (*squared*)



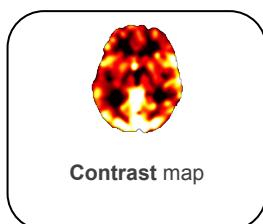
3dMEMA\_result+tlrc.BRIK[[0]]  
[from contrast & stat maps]



spmT\_0001.nii

tstat1.nii.gz  
zstat1.nii.gz

3dMEMA\_result+tlrc.BRIK[[1]]

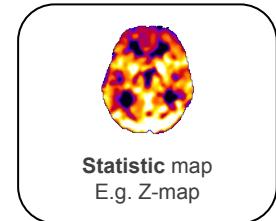


con\_0001.nii

cope1.nii

3dMEMA\_result+tlrc.BRIK[[0]]

# IBMA on Z maps



- Fisher's

$$-2 \sum_k \log P_k \sim \chi^2_{2k}$$

- Sum of -log P-values (from T/Z's converted to P's)

- Stouffer's

$$\sqrt{K} \times \frac{1}{K} \sum_k Z_k \sim \mathcal{N}(0, 1)$$

- Average Z, rescaled to N(0,1)

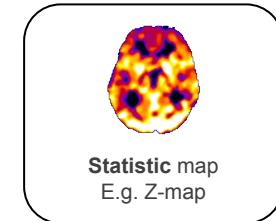
- “Stouffer's Random Effects (RFX)”

$$\sqrt{K} \times \frac{1}{K} \sum_k Z_k \sim \mathcal{N}(0, \sigma_{RFX}^2)$$

- Submit Z's to one-sample t-test

(Slide adapted from Thomas Nichols, OHBM 2015)

# IBMA on Z maps + N



+ N

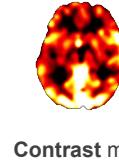
- Weighted Stouffer's

$$\sum_k w_k Z_k \sim \mathcal{N}(0, 1), \quad w_k \propto \sqrt{N_k}$$

- Z's from bigger studies get bigger weights

(Slide adapted from Thomas Nichols, OHBM 2015)

# IBMA on Contrast maps

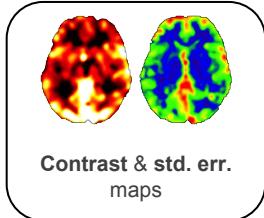


- Random Effects (RFX) GLM

$$\frac{1}{K} \sum_k c\hat{\beta}_k \sim \mathcal{N}(0, \sigma_{\text{RFX}}^2)$$

- Analyze per-study contrasts as “data”

## Contrast + standard error maps



- Fixed-Effects (FFX) GLM

$$\frac{1}{K} \sum_k \hat{\theta}_k \sim \mathcal{N}(0, \sum_k \sigma_{\text{FFX},k}^2 / K^2)$$

- *Don't estimate variance, just take from first level*

(Slide adapted from Thomas Nichols, OHBM 2015)

# Implementations

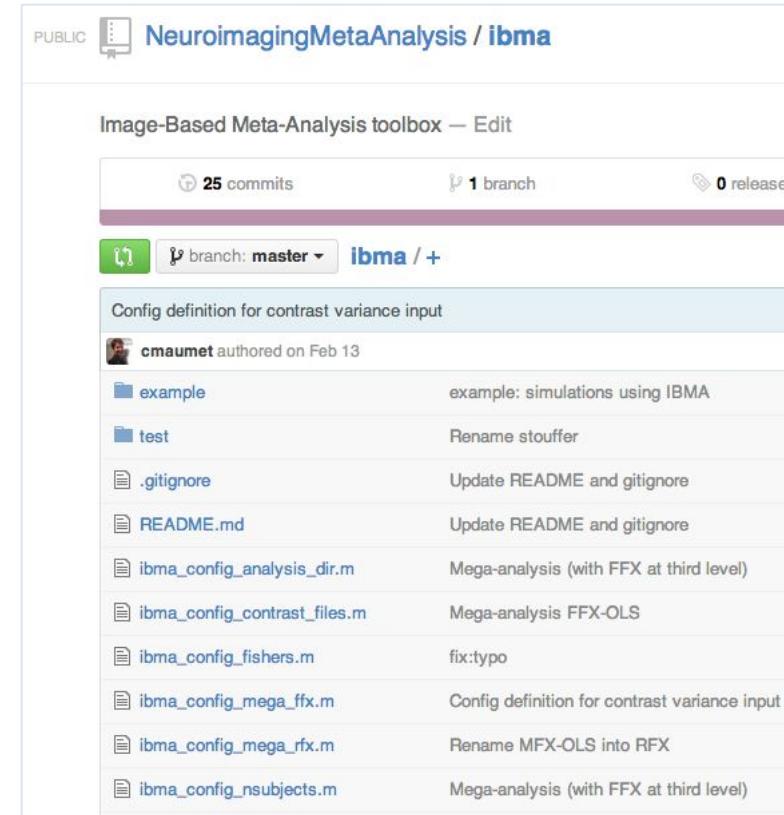
- Not all of these options are easily used

Meta-Analysis Method	Inputs	Neuroimaging Implementation
'Gold Standard' MFX	Con's + SE's	FSL's FEAT SPM <code>spm_mfx</code> AFNI 3dMEMA
RFX GLM	Con's	FSL, SPM, AFNI, etc...
Stouffer's RFX	Z's	
FFX GLM	Con's +SE's	
Fisher's	Z's	n/a
Stouffer's	Z's	
Stouffer's Weighted	Z's + N's	

(Slide from Thomas Nichols, OHBM 2015)

# Self Promotion Alert: IBMA toolbox

- SPM Extension
- Still in beta!
  - But welcome all feedback
- Available on GitHub



<https://github.com/NeuroimagingMetaAnalysis/ibma>

a

# Meta-analysis of 21 pain studies

- Results
  - GLM methods similar
  - Z-based methods similar
    - But FFX Z methods more sensitive (as expected)

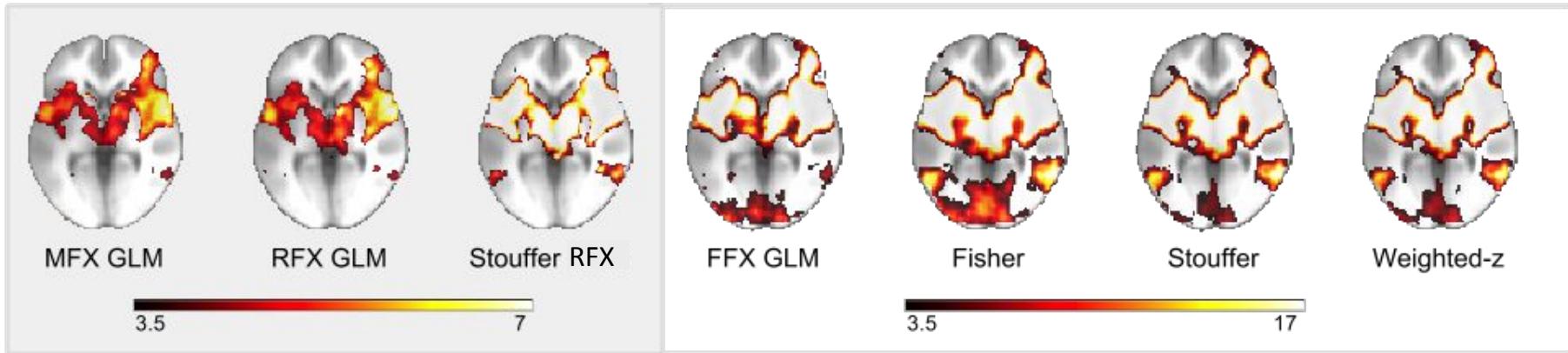
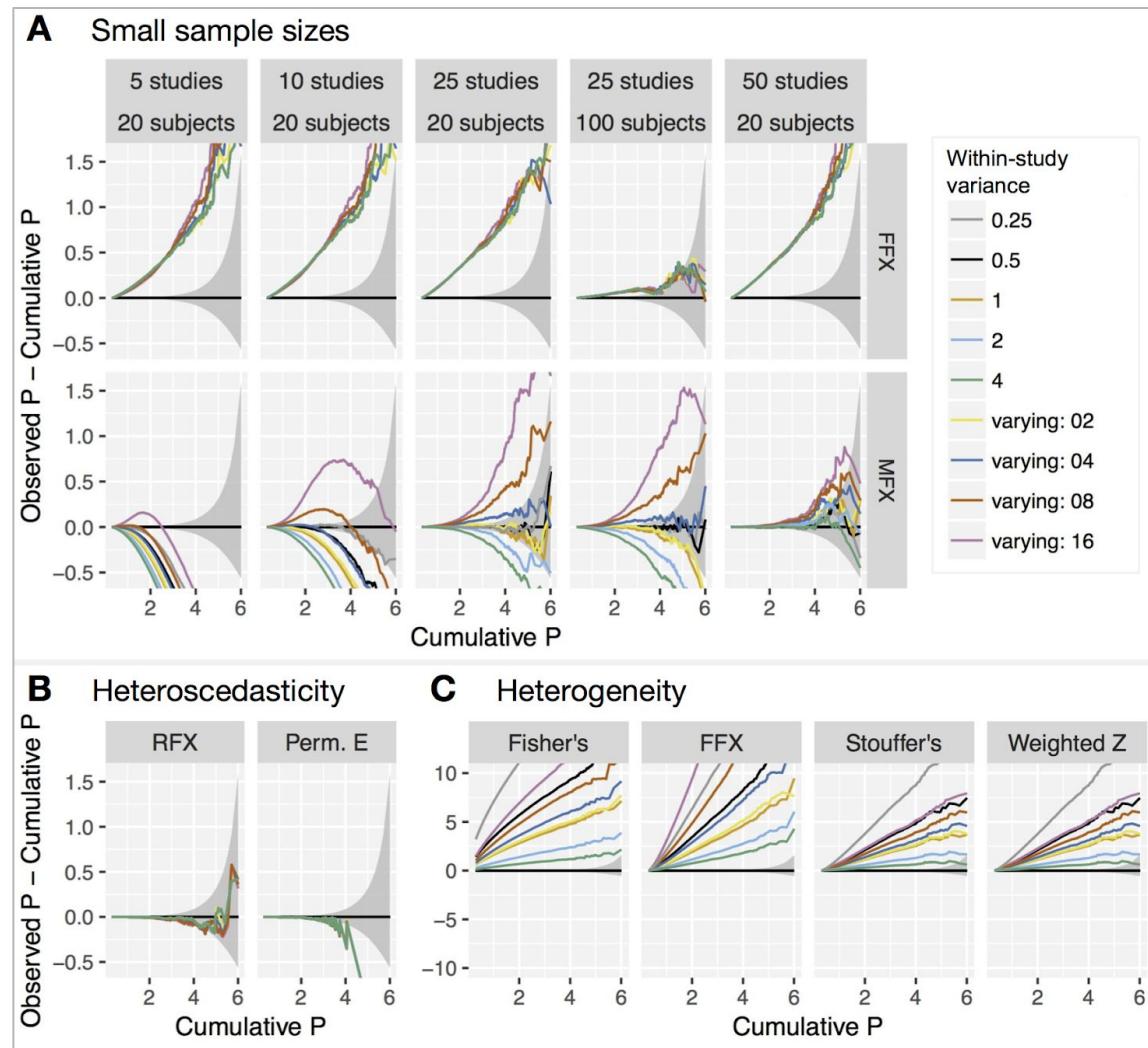


Fig. 1: Result of a meta-analysis of 21 pain studies for 4 fixed-effects (FFX GLM, Fisher, Stouffer, weighted-z) and 2 random-effects (RFX GLM, Stouffer MFX) meta-analytic approaches compared to the reference (MFX GLM) at a threshold of  $p < 0.05$  FDR corrected.

Data: Tracey pain group, FMRIB, Oxford.

# Self Promotion Alert: Robustness of the meta-analytic estimators

Poster 2653

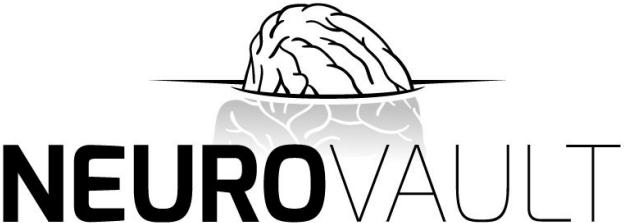


# How to publish your statistic maps?

# Share your statistic maps

NeuroVault Collections ▾ FAQ Give feedback

Log in



A public repository of unthresholded statistical maps, parcellations, and atlases of the human brain

What is it?

A place where researchers can publicly store and share unthresholded statistical maps, parcellations, and atlases produced by MRI and PET studies.

Why use it?

- Interactive visualization
- A permanent URL
- Publicly shareable
- Improves meta-analyses

Get started and upload an image!

Supported by



<http://neurovault.org>

# Share your statistic maps

NeuroVault Collections FAQ Give feedback Log in

## A Correspondence between Individual Differences in the Brain's Intrinsic Functional Architecture and the Content and Form of Self-Generated Thoughts

Contributed by ChrisFilipGorgolewski

Krzysztof J. Gorgolewski, Dan Lurie, Sebastian Urchs, Judy A. Kipping, R. Cameron Craddock, Michael P. Milham, Daniel S. Margulies, Jonathan Smallwood

[Link to the paper](#)

[3D View](#)

File View Options

x y z 0  
0 -18 18 0

Images Details

Show 7 entries Search:

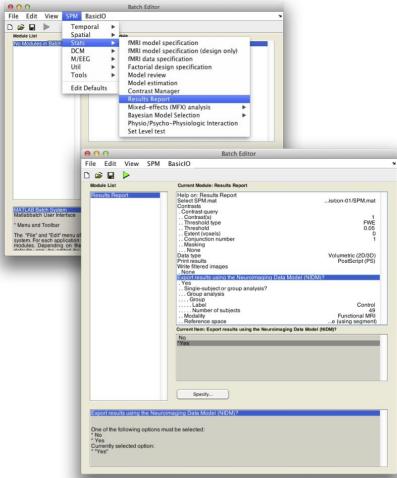
View	ID	Name	Type
	25	fALFF: Past	Z map
	26	fALFF: Future	Z map
	27	fALFF: Past > Future	Z map
	28	fALFF: Positive	Z map
	29	fALFF: Negative	T map
	30	fALFF: Positive > Negative	Z map
	31	fALFF: Social Cognition	Z map

Showing 1 to 7 of 30 entries [First](#) [Previous](#) [Next](#) [Last](#)

<http://neurovault.org>

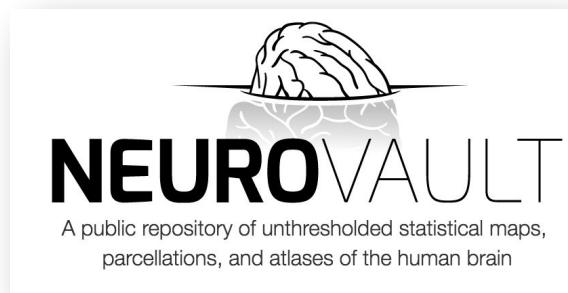
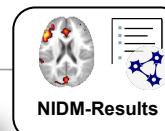
# From SPM & FSL

SPM



Name	Size	Kind
beta_0001.nii	2.4 MB	NIFTI
con_0001.nii	2.4 MB	NIFTI
mask.nii	593 KB	NIFTI
nidm_001.nidm.zip	4.1 MB	ZIP archive
ResMS.nii	4.7 MB	NIFTI
RPV.nii	4.7 MB	NIFTI
SPM.mat	484 KB	MATLAB Data
spmT_0001.nii	2.4 MB	NIFTI

Name	Size	Kind
design.con	126 bytes	Document
design.fsf	20 kB	Document
design.grp	131 bytes	Document
design.icon	28 bytes	Document
cope1.feat	--	Folder
cope2.feat	--	Folder
inputreg	--	Folder
logs	--	Folder
bg_image.nii.gz	3.4 MB	gzip compressed archive
mask.nii.gz	15 kB	gzip compressed archive
mean_image.nii.gz	2.8 MB	gzip compressed archive
mean_center.nii.gz	2.8 MB	gzip compressed archive
report.html	10 kB	HTML document
report_reff.html	55 kB	HTML document
report_res.html	17 kB	HTML document
report_stats.html	495 bytes	HTML document
report.html	922 bytes	HTML document
design.mat	732 bytes	MATLAB Data
design_cov.ppm	6 kB	OpenOffice.app Document
design.ppm	219 kB	OpenOffice.app Document
design_cov.png	106 bytes	Portable Network Graphics Image
design.png	1 kB	Portable Network Graphics Image
fsl_ds107_group.nidm.zip	13.6 MB	ZIP archive



```
$ nidmfsl fsl_ds107_group 49 -g Control
```

<http://nidm.nidash.org/getting-started/>

# Conclusions

- When data available, **Image-Based** preferred to **Coordinate-Based** meta-analysis

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- Few tools for Z-based IBMA, but underway...

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- **In practice**, it is **difficult** to use the gold standard **Mixed-Effects GLM**
- When only contrast estimates are available, **RFX GLM** is a practical & valid approach
- Few tools for Z-based IBMA, but underway...
- Data sharing tools: **NeuroVault**, **NIDM-Results**

# Thank you!

This work is supported by

