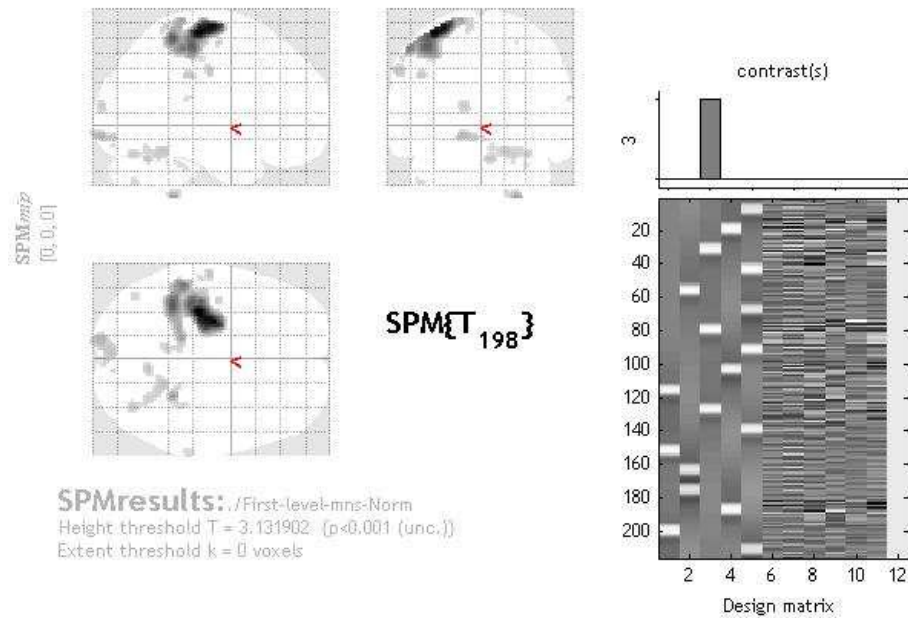
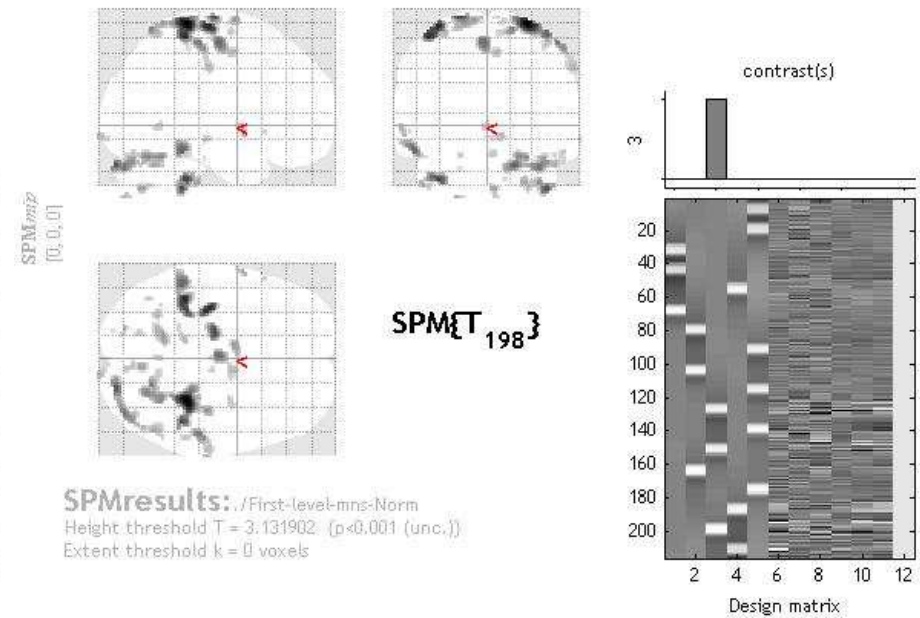


Laterality index in brain imaging research

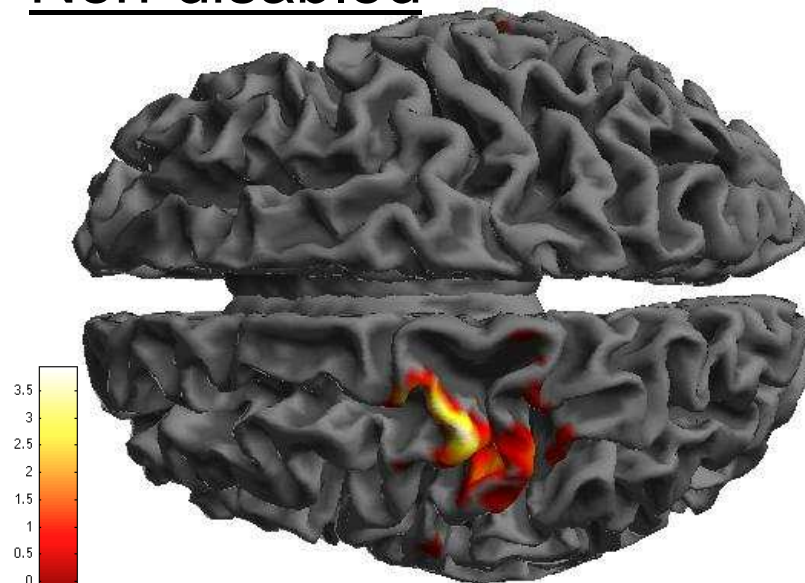
ER - All Sessions



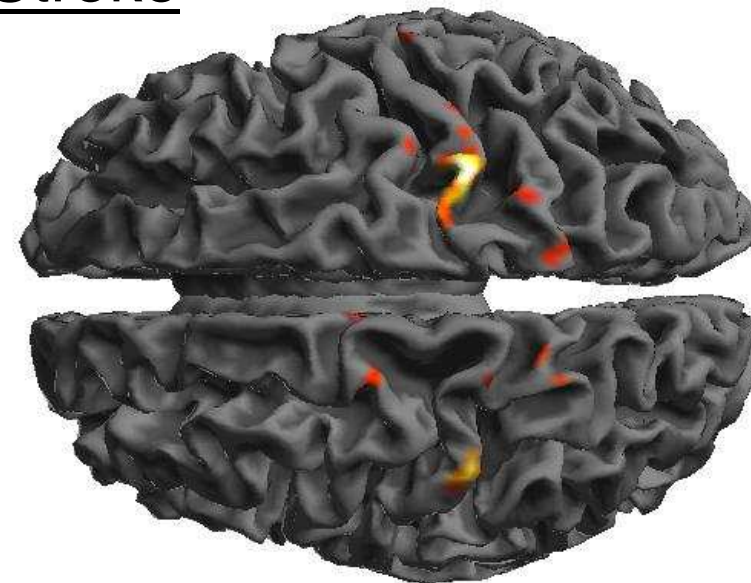
ER - All Sessions



Non-disabled



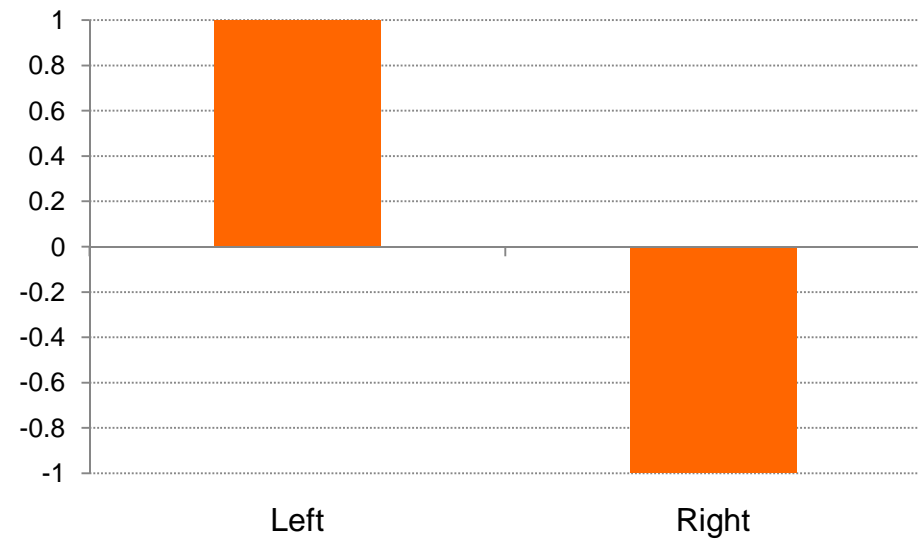
Stroke



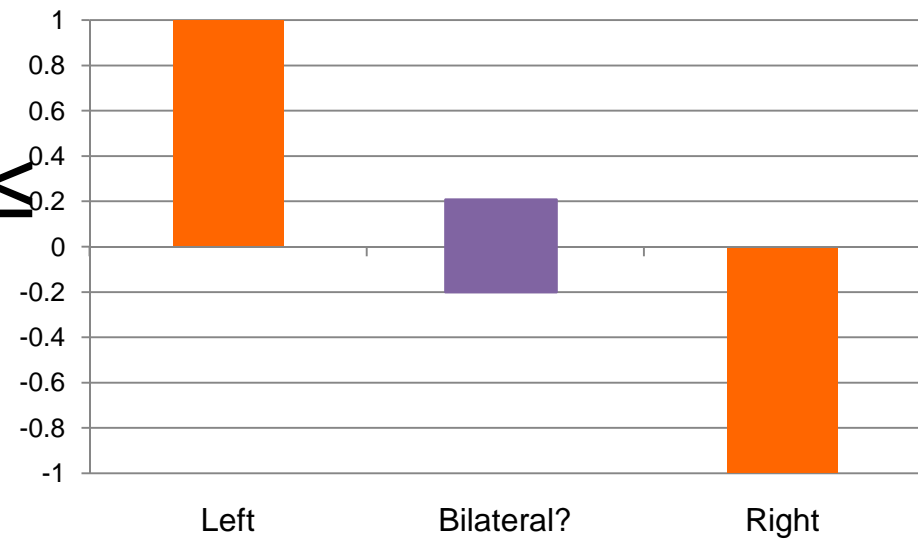
$$LI = \frac{\text{Left} - \text{Right}}{\text{Left} + \text{Right}}$$

$LI > 0 = \text{Left}$

$LI < 0 = \text{Right}$

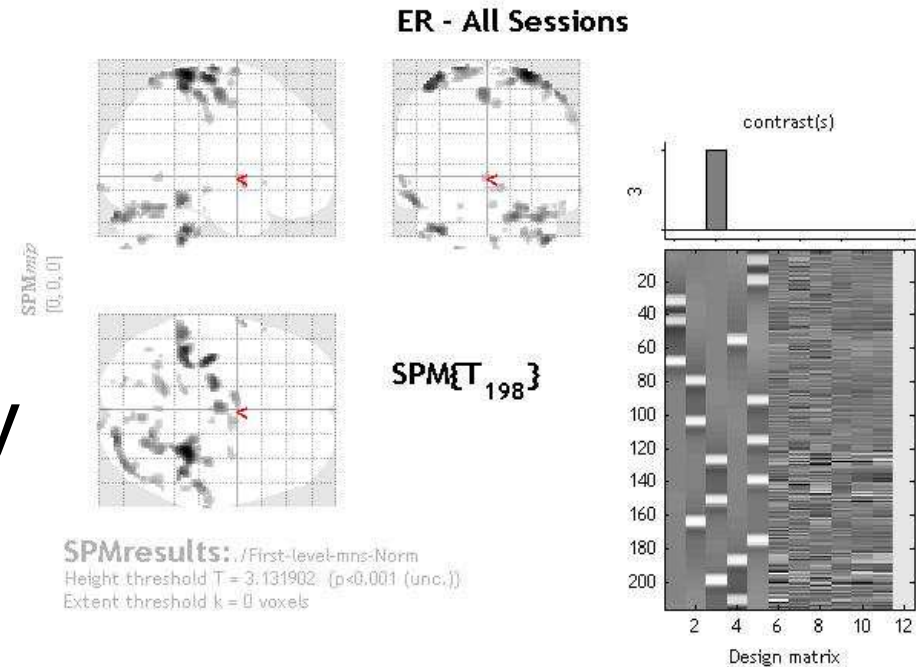


$$|LI| \leq 0.2 = \text{Bias} \leq$$



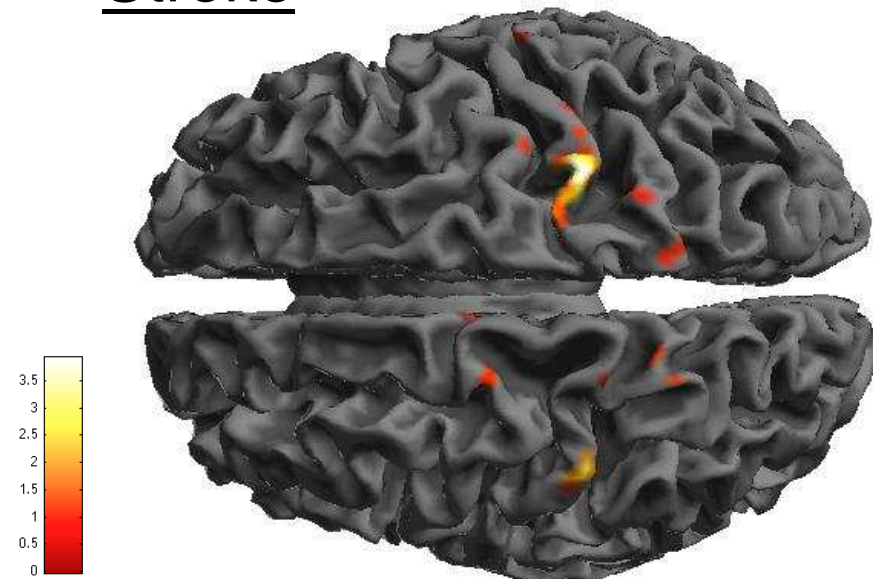
LI affected by:

1. Measure of activity
 - A. Extent
 - B. Magnitude



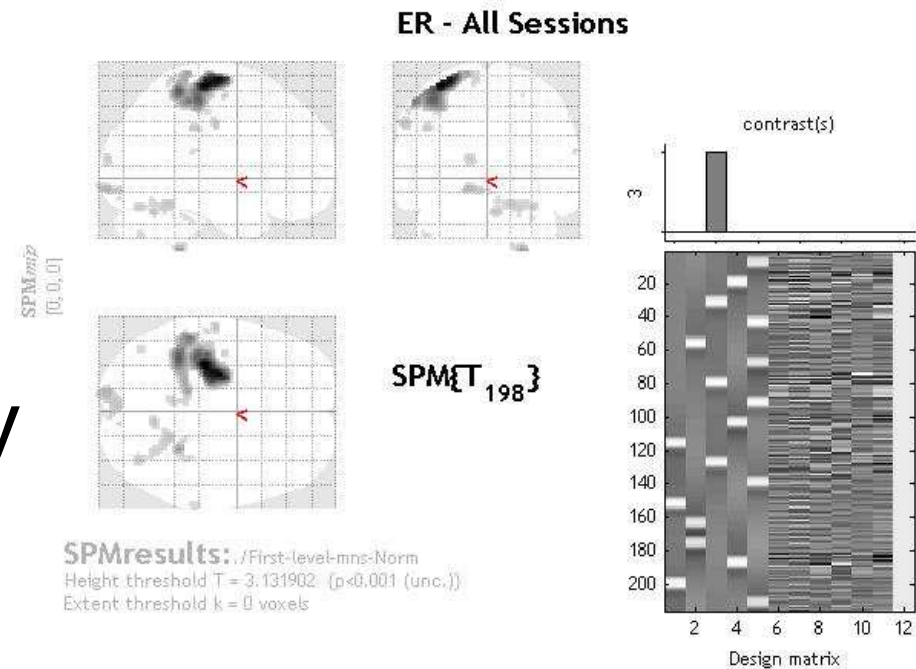
Stroke

2. Choice of ROI



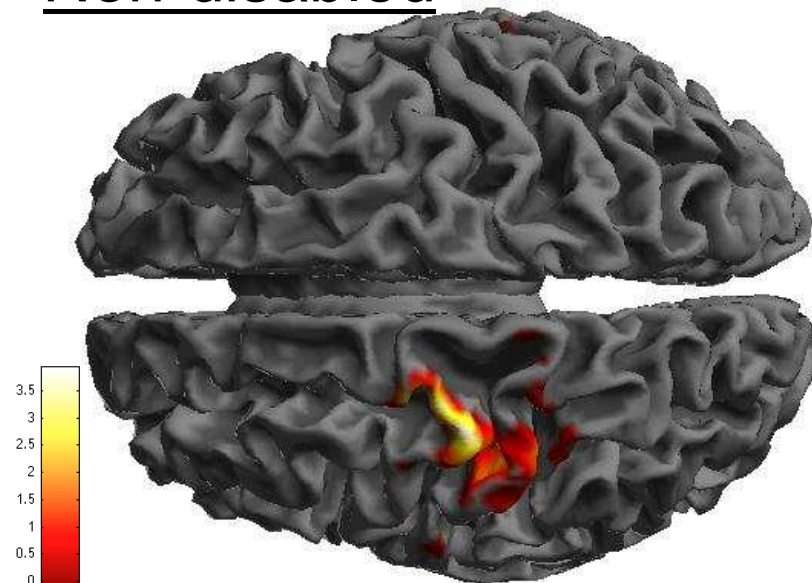
LI affected by:

1. Measure of activity
 - A. Extent
 - B. Magnitude



Non-disabled

2. Choice of ROI



A. Extent - # voxels

B. Magnitude - % signal change

A. Extent - # voxels *at variable thresholds*

B. Magnitude - % signal change *of most active*

A. LI based on extent of activity:

Wilke, 2006, 2007

LI-toolbox **SPM2** **SPM5** **SPM8**

Summary: Allows assessment of laterality effects in imaging data using various thresholding options. Among other features, regionally-restricted analyses are possible and a bootstrapping approach allows to assess data homogeneity to reduce the effect of outliers. The toolbox can now be scripted, allowing for unattended analyses. Matlab from version 6.5.1 on is required.

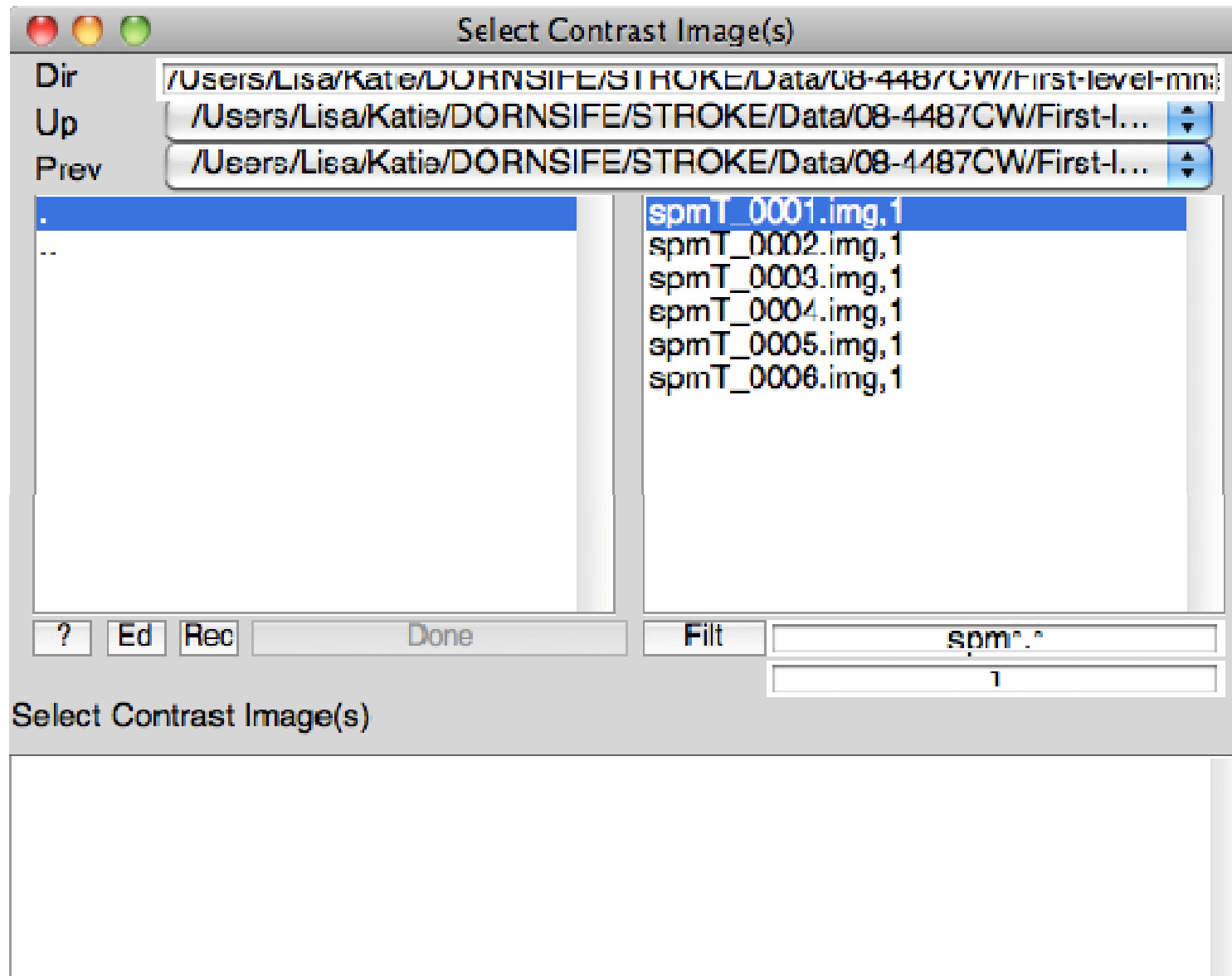
Author: [Marko Wilke](#)

URL: [Contact Marko by email.](#)

file:///Applications/MATLAB_R2010a/spm8/toolbox/LI/man/index.html

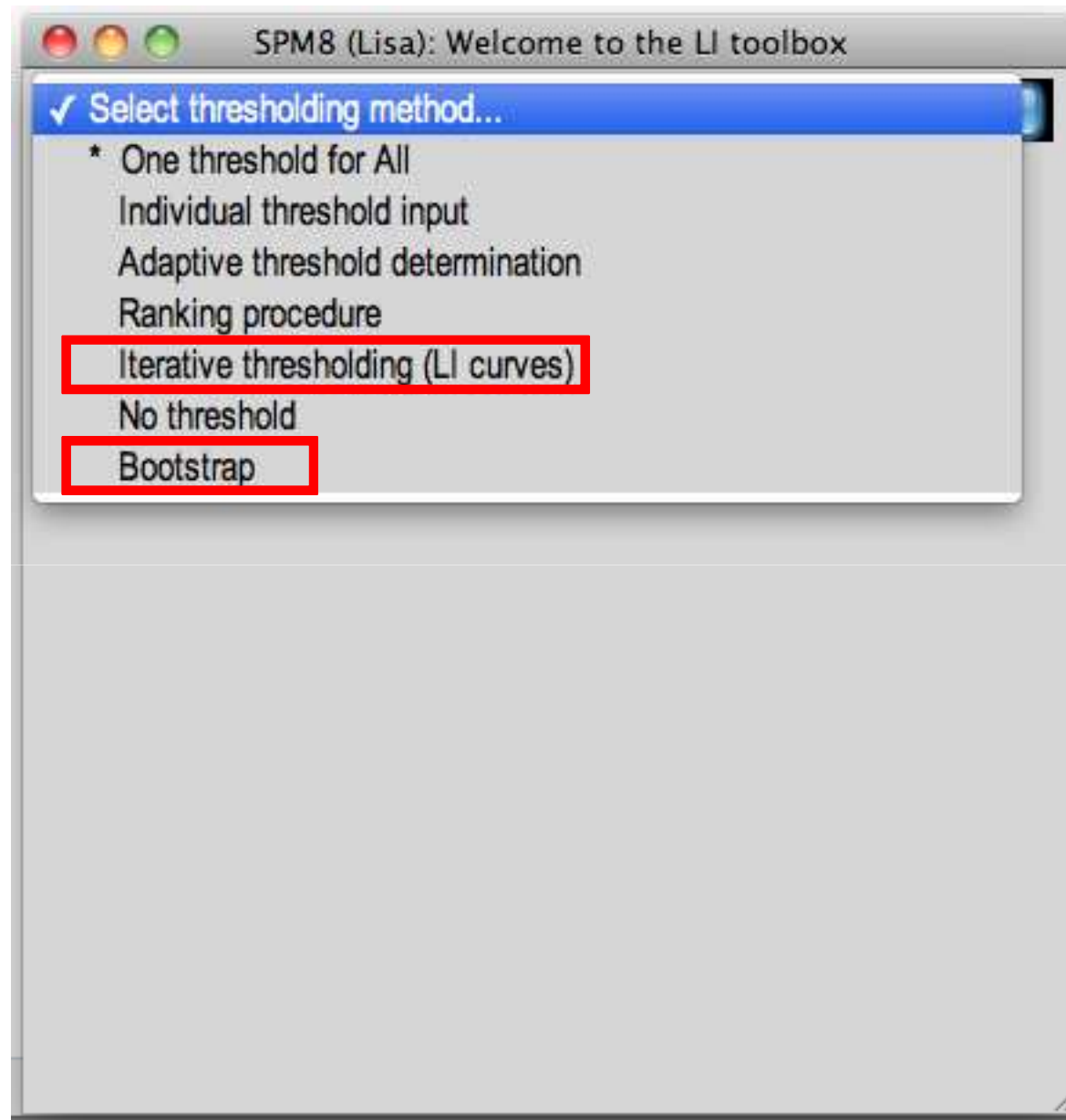
1. Select contrast image

2/16



2. Select threshold method

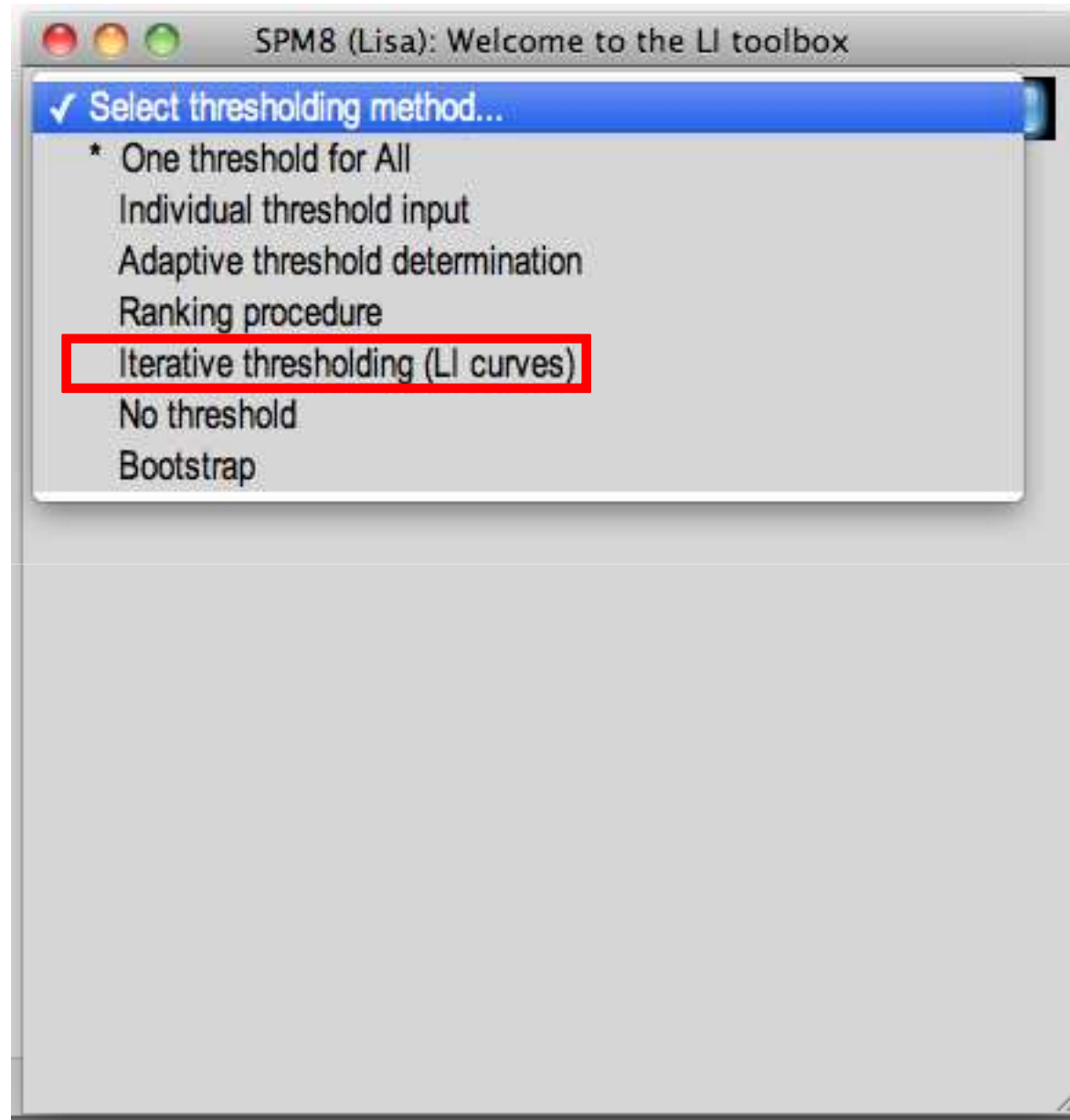
3/16



LI based on extent = dependen

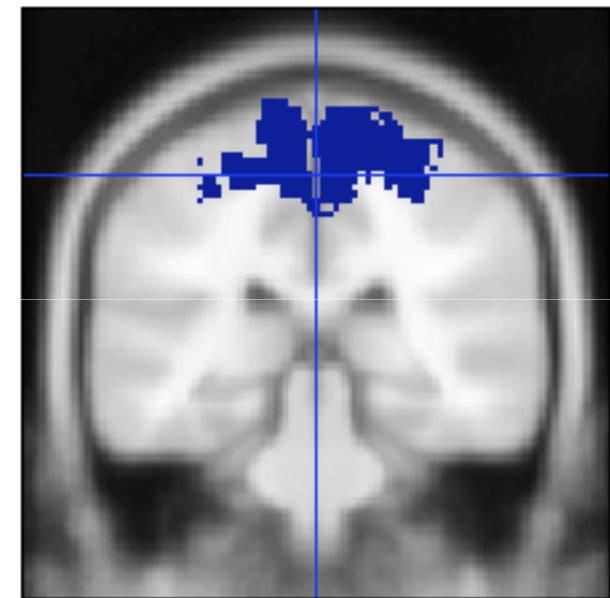
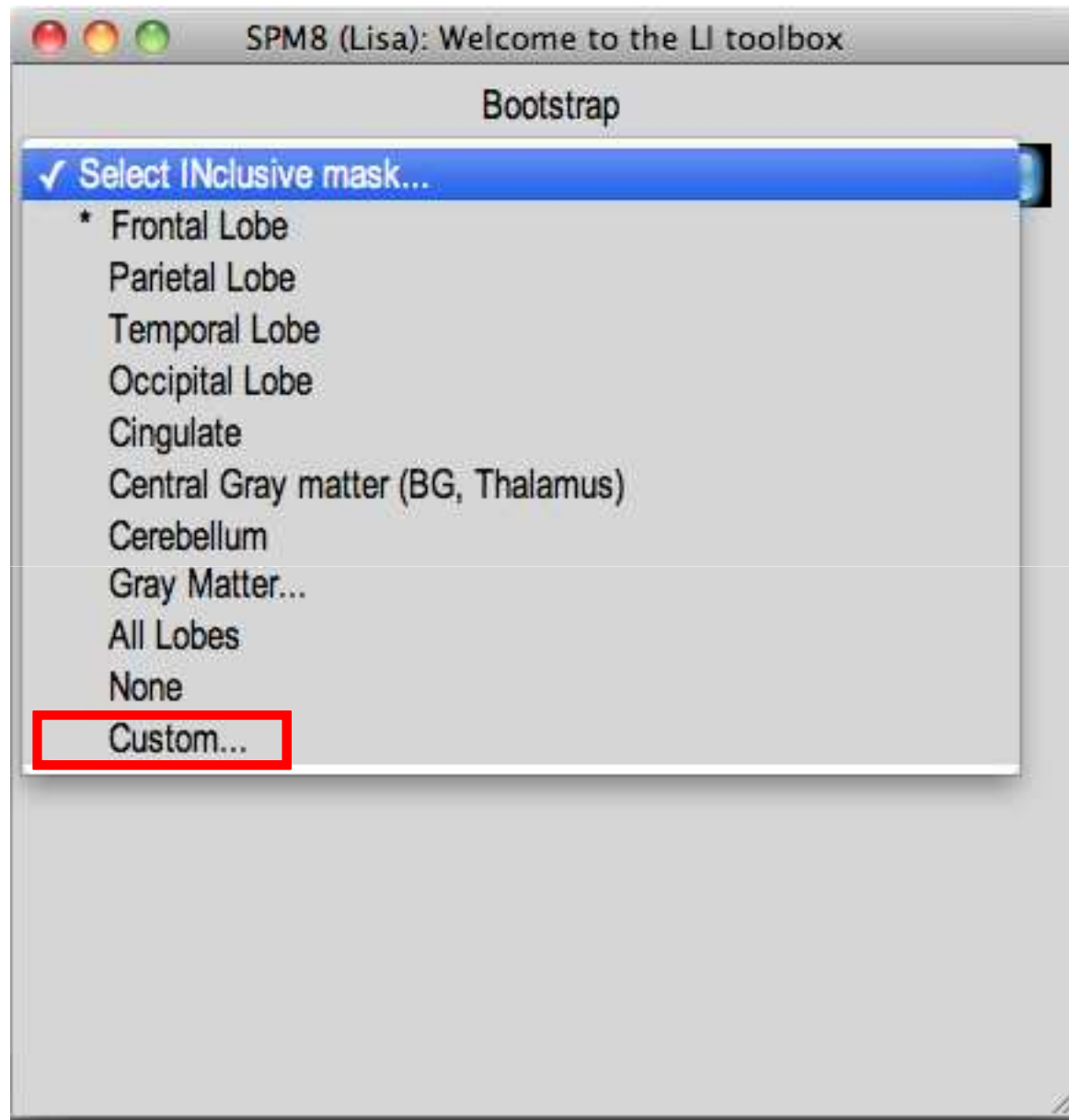
2. Select iterative threshold method

4/16



3. Select inclusive mask

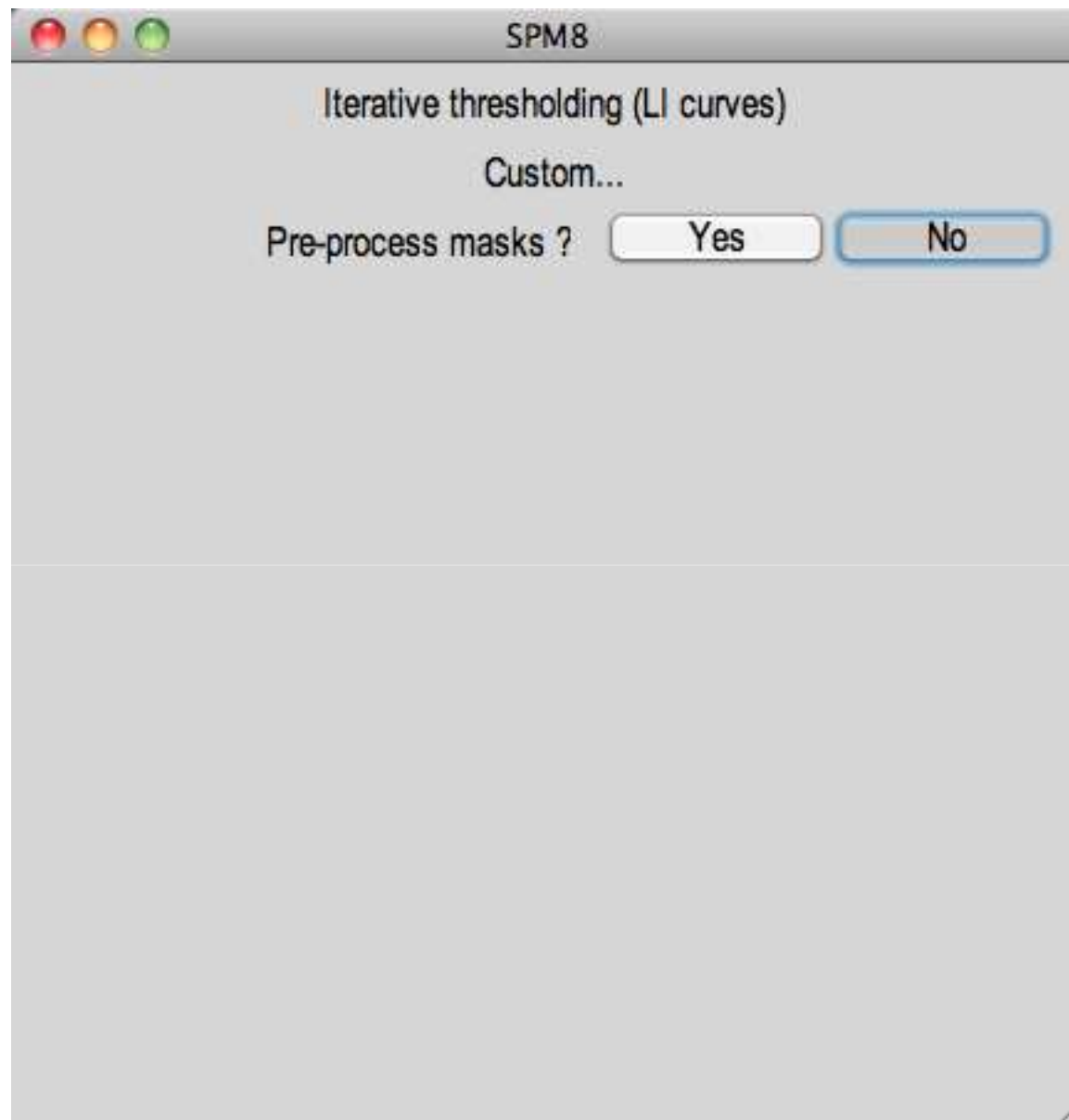
5/16



Primary motor cortex
BA 4a + 4p

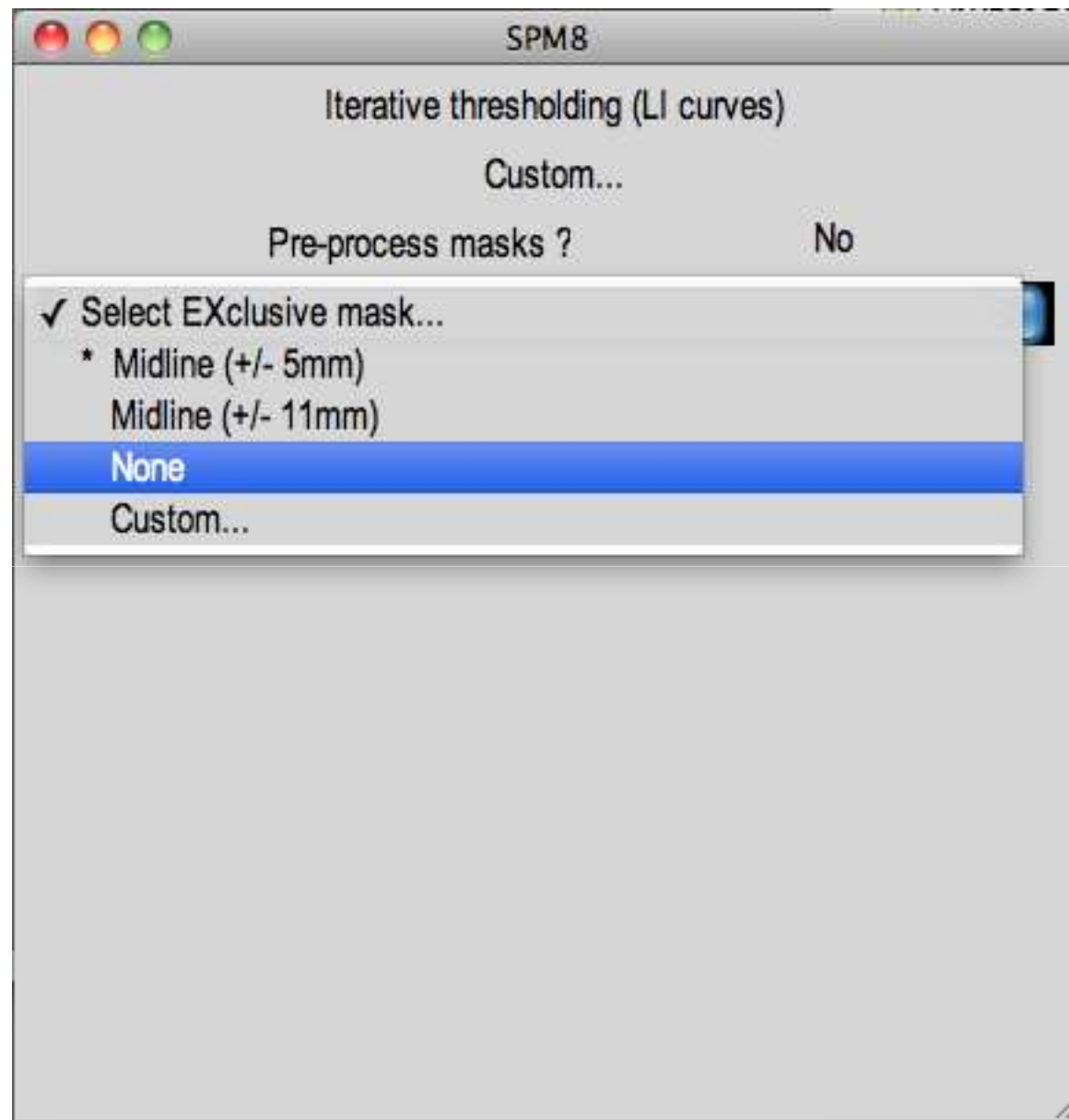
4. Pre-process masks?

6/16



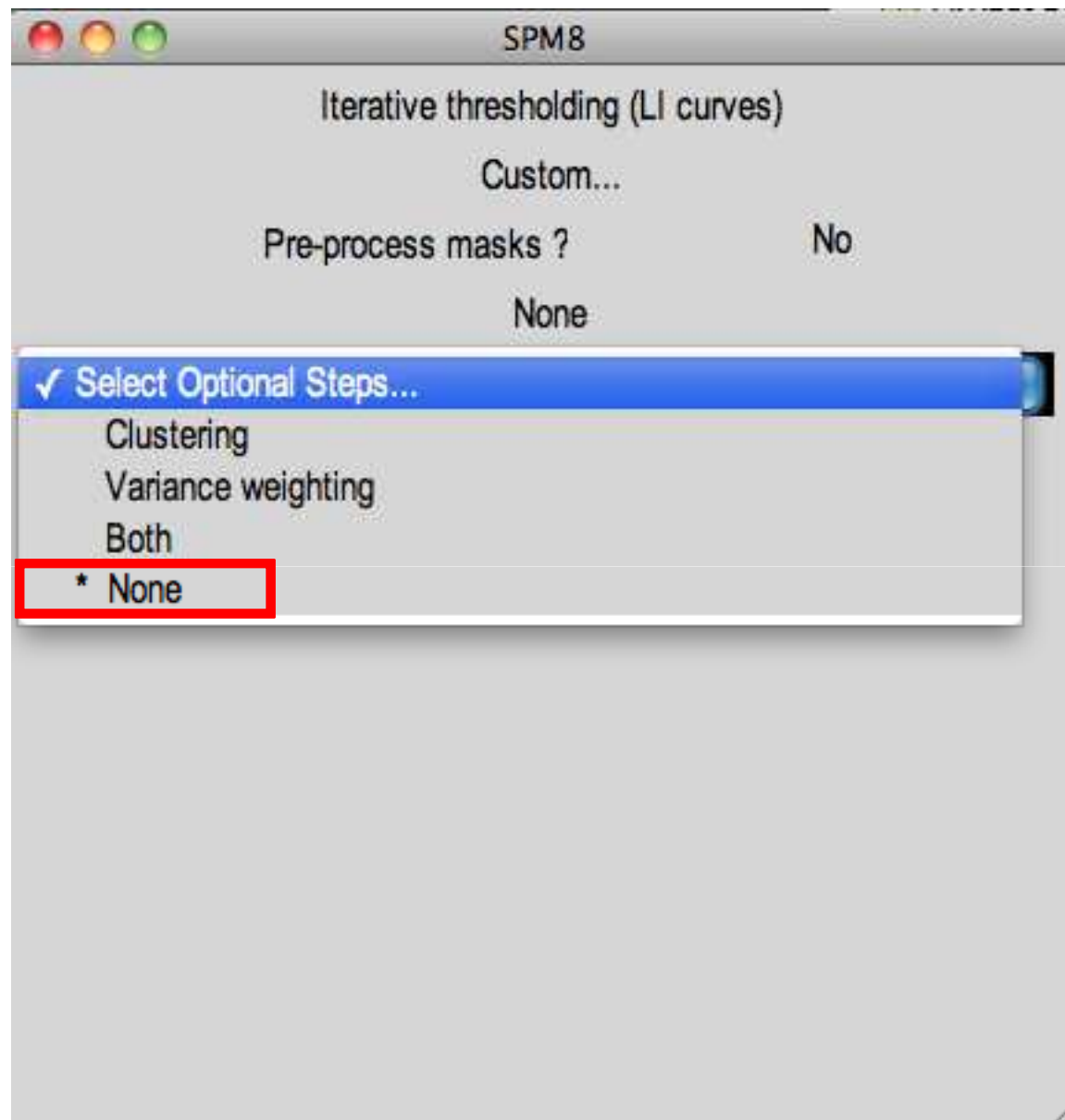
5. Select exclusive mask

7/16



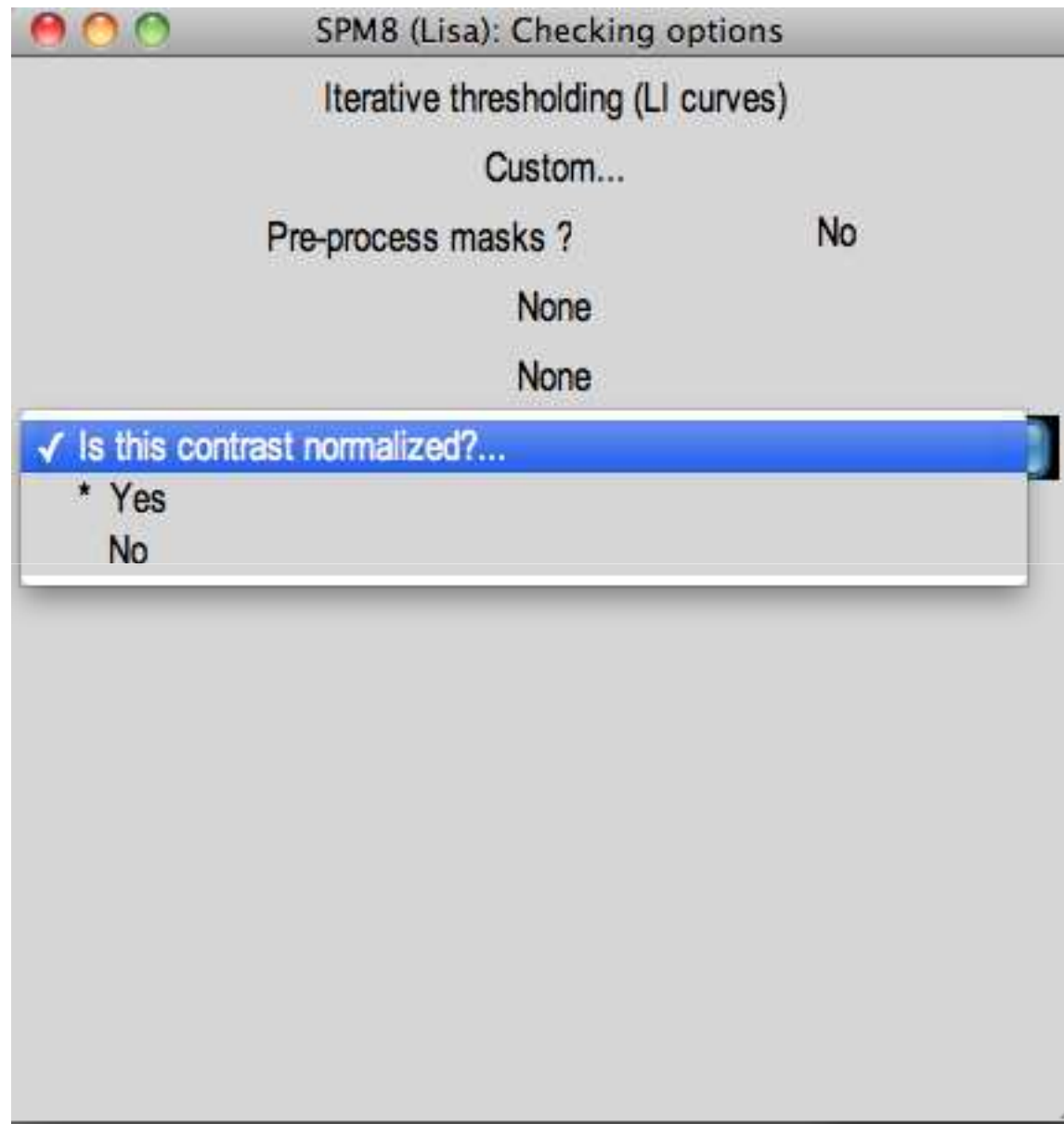
6. Optional steps

8/16



7. Is the contrast normalized?

9/16



Iterative thresholding output:

10/16

- LI report
- LI curves

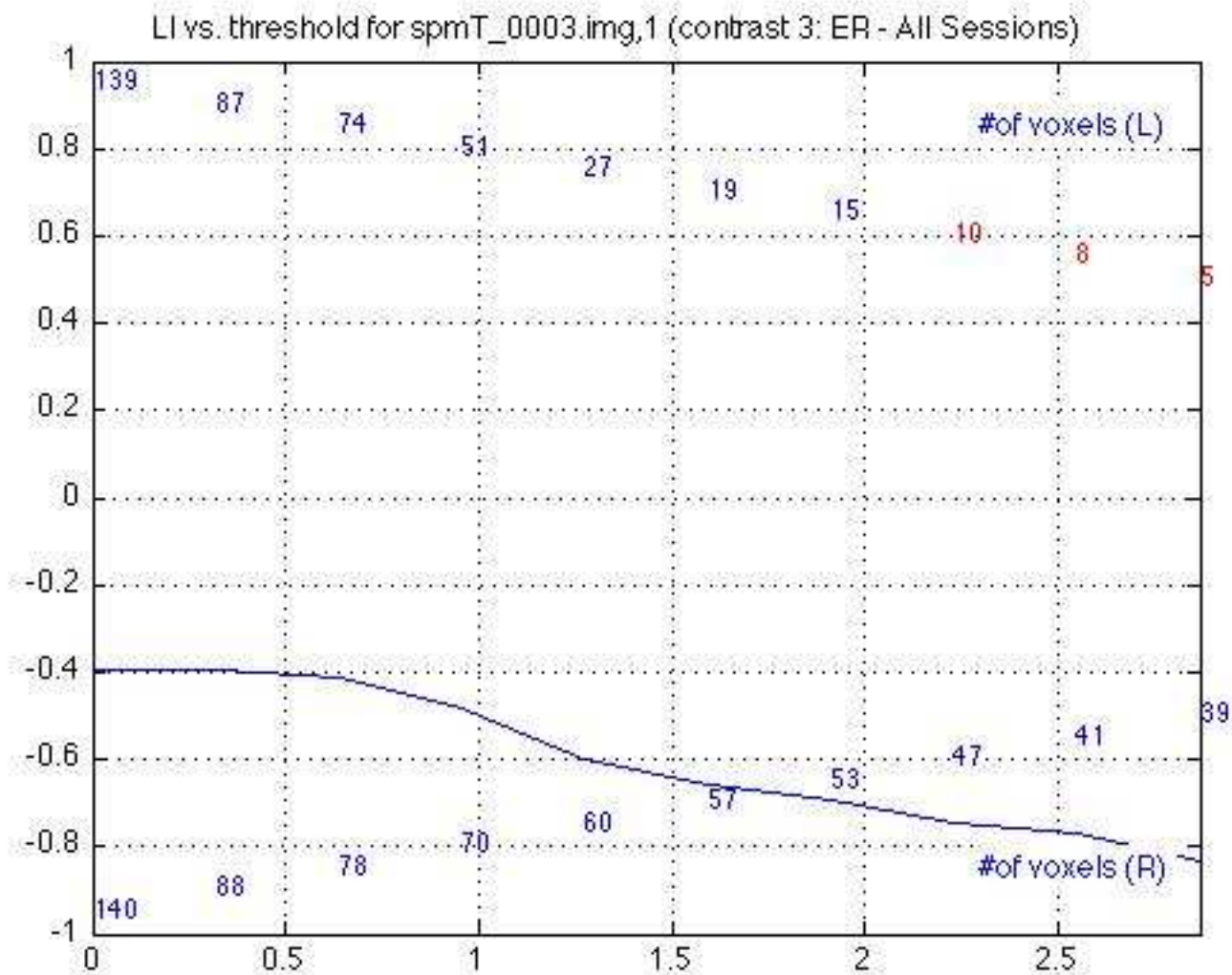
LI = [-39 to -.83]

- *weighted by voxel value
- *you can specify voxel count instead
- *overall LI results are similar
- *take the mean to report

Iterative thresholding output:

11/16

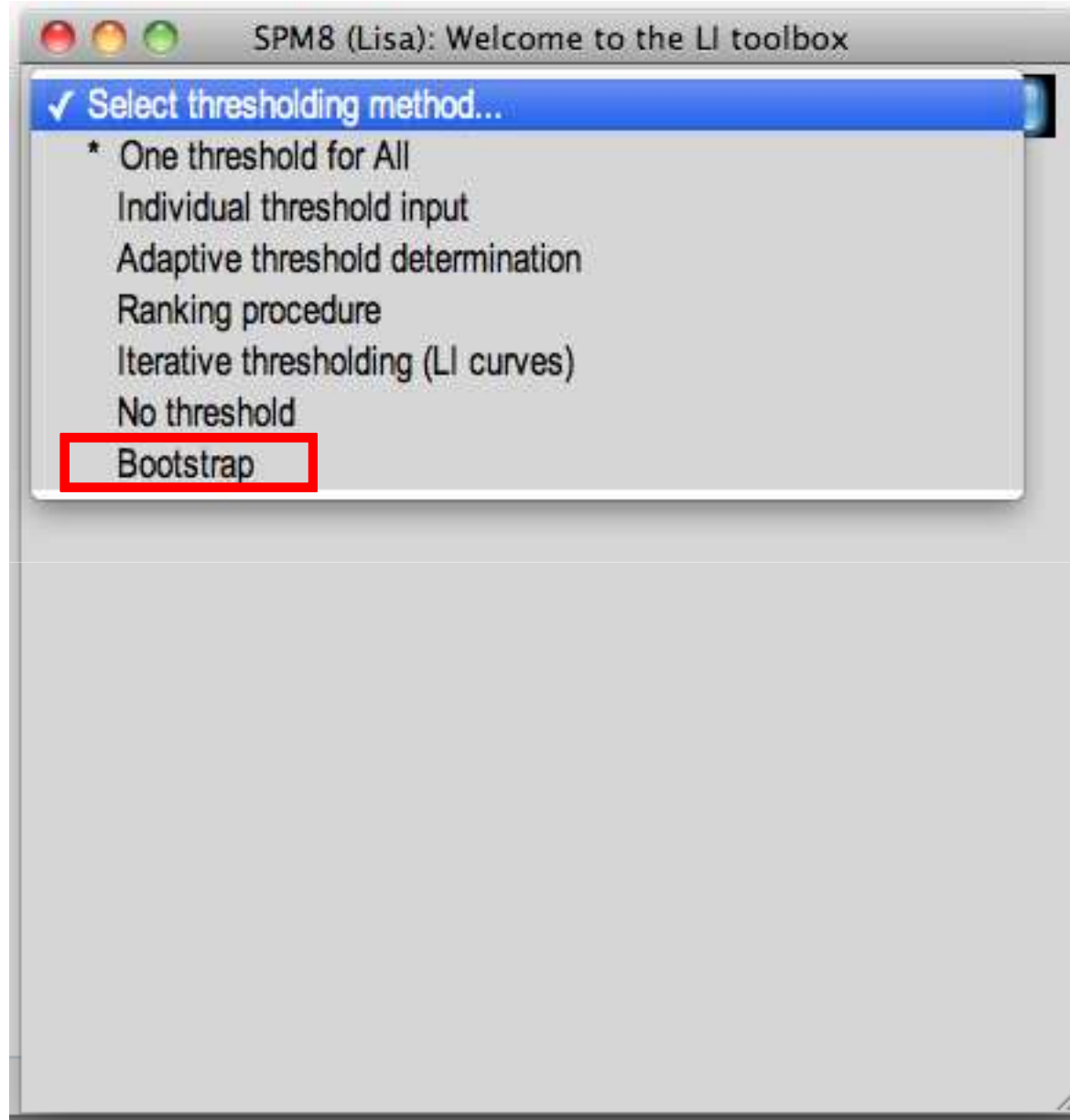
- LI report
- LI curves



Incl: 11-BA-4an-Bilateral roi mat nii 1 Excl: none Clust: no VarWeight: no

Back to 2. Select bootstrap threshold method:

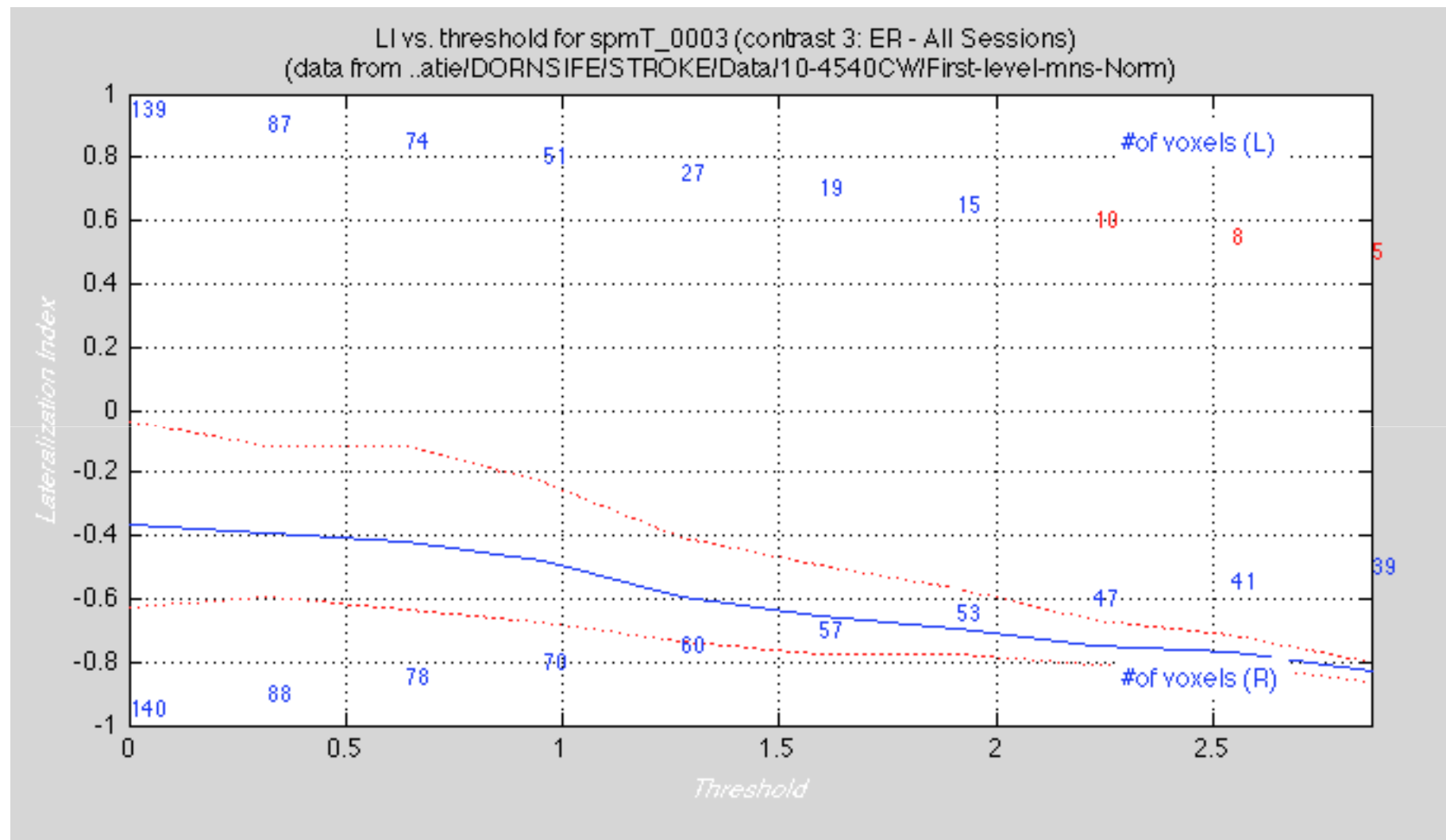
12/16



Wilke, 2006

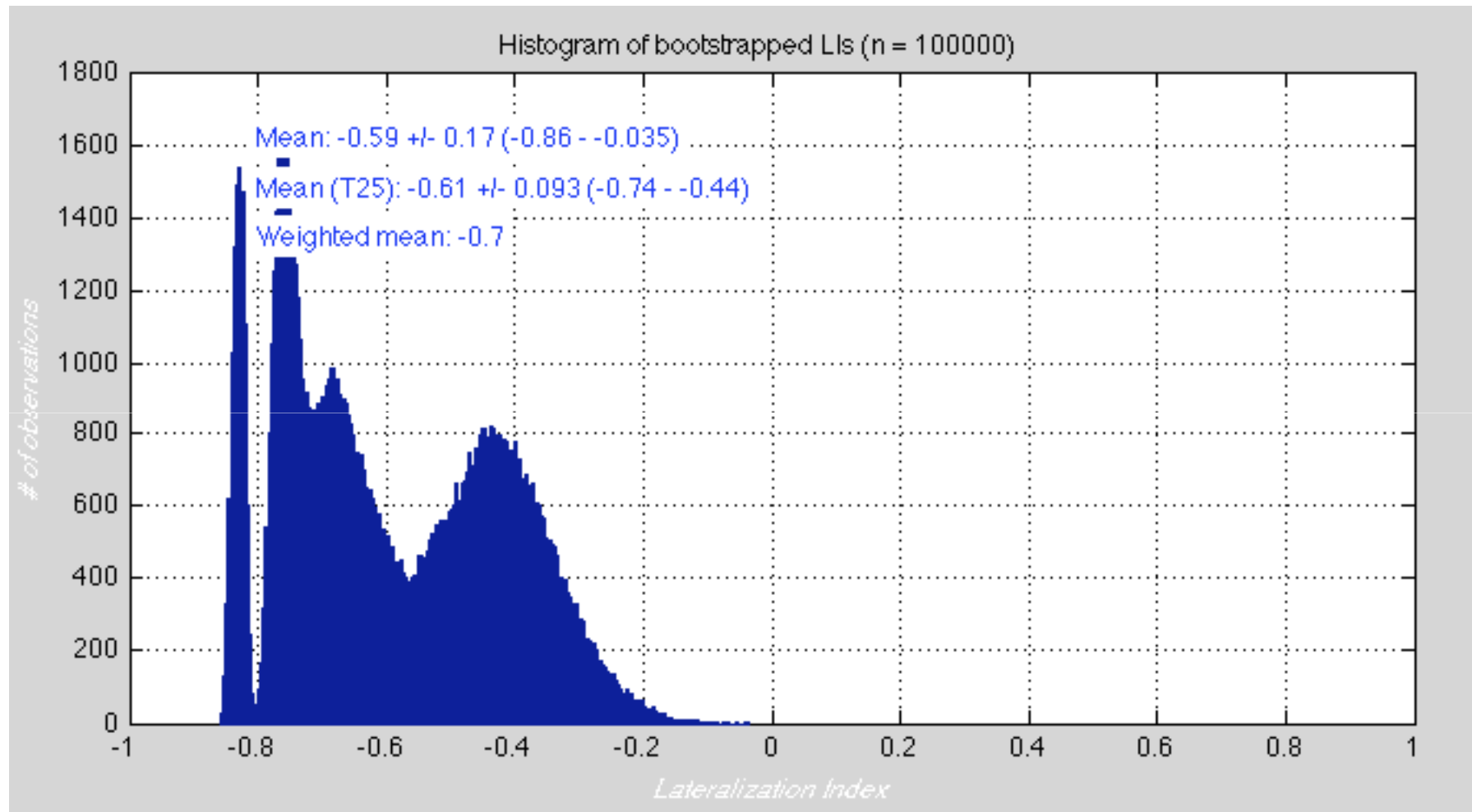
Bootstrap output:

13/16

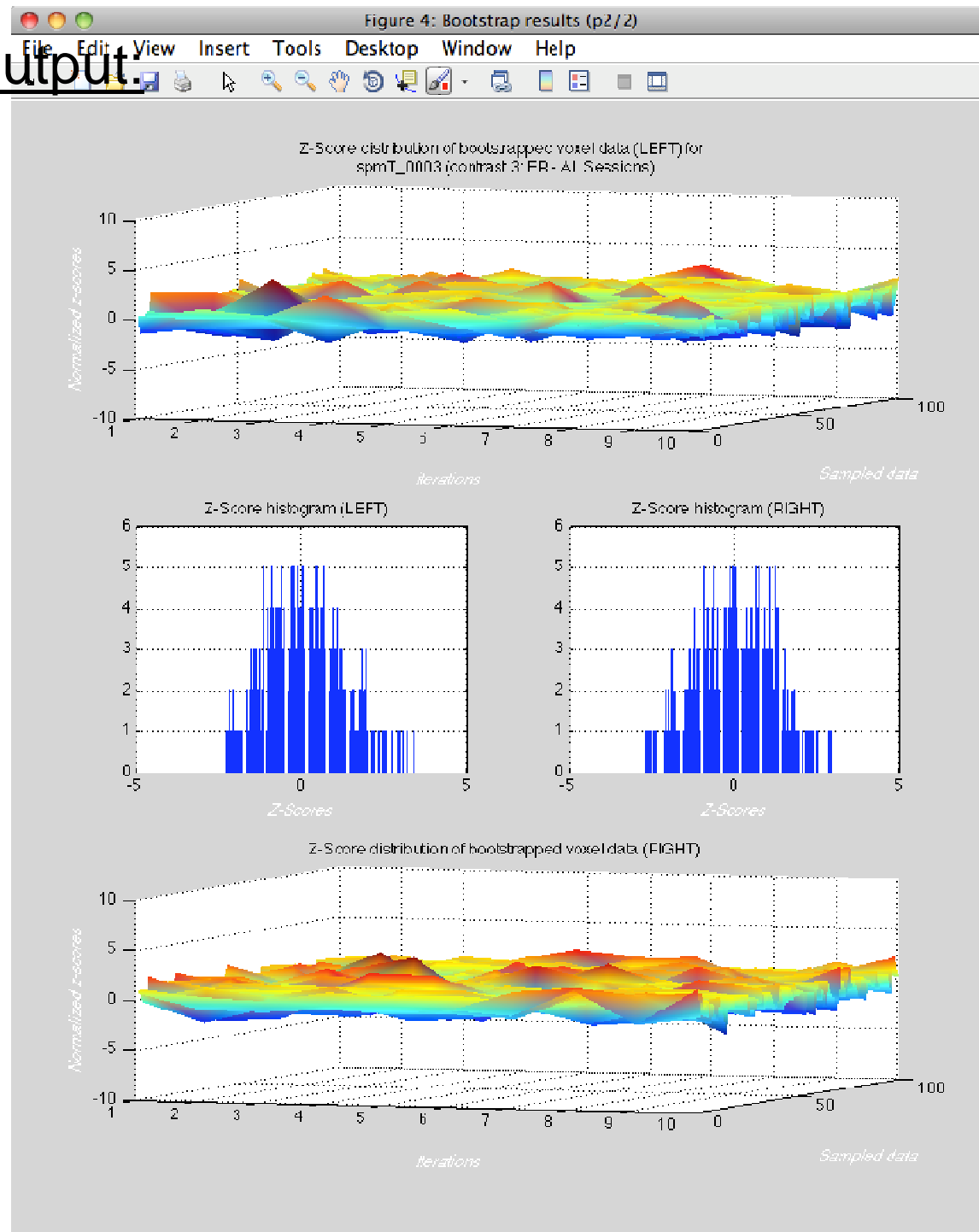


Bootstrap output:

14/16



Bootstrap output



Bootstrap output:

16/16

- Overall bootstrap mean LI = $-.59$
 - results from all thresholds, along x-axis
 - trimmed and weighted
- Weighted mean LI = $-.70$
 - based on results from all thresholds along LI curve x-axis, 'v'
 - e.g. voxels that survive higher thresholds likely correlate more with 'v'
- Trimmed mean LI = $-.61$
 - based on the trimmed means at each threshold, punishes 'v'
- Which to use depends on your question, region, 'v'

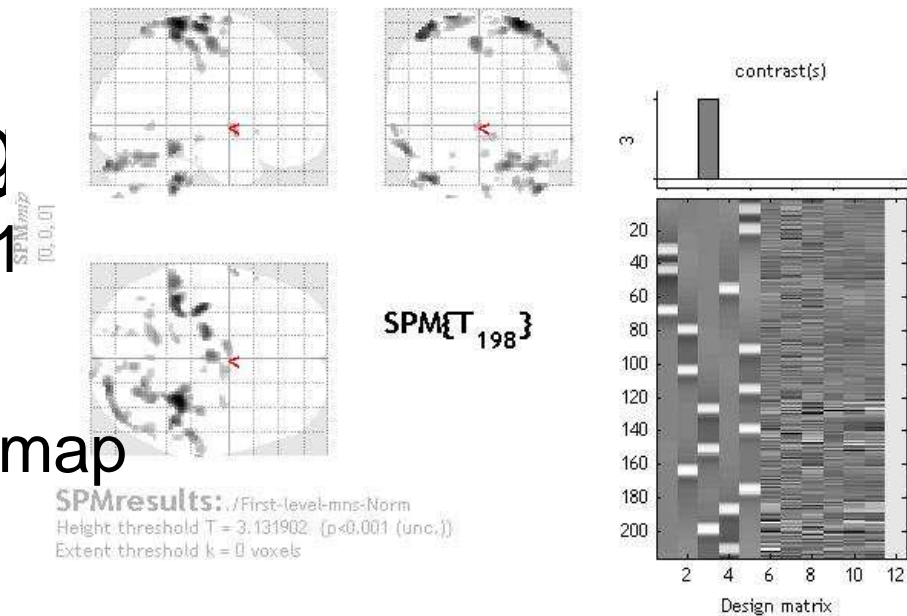
B. LI based on mag

Fernandez, 2001

First, get single subject T-map
at $p < .001$ uncorrected

ER - All Sessions

1/5



Statistics: *p-values adjusted for search volume*

set-level		cluster-level					peak-level					mm mm mm		
p	c	$p_{FWECorr}$	$q_{FDRcorr}$	k_c	p_{uncorr}	$p_{FWECorr}$	$q_{FDRcorr}$	T	(Z_e)	p_{uncorr}				
0.000	39	0.003	0.007	272	0.000	0.000	0.000	8.14	7.55	0.000	30	-36	72	
						0.002	0.001	5.72	5.50	0.000	44	-38	64	
						0.143	0.027	4.66	4.59	0.000	36	-22	68	
		0.592	0.209	54	0.054	0.000	0.000	7.05	6.65	0.000	-36	-20	70	
		0.199	0.110	97	0.013	0.000	0.000	7.04	6.64	0.000	-42	-38	62	
		0.768	0.311	41	0.088	0.000	0.000	6.41	6.10	0.000	-34	-36	-38	
		0.540	0.209	58	0.047	0.000	0.000	6.37	6.07	0.000	-20	-16	76	
		0.553	0.209	57	0.048	0.000	0.000	6.24	5.95	0.000	44	-40	-44	
						0.073	0.015	4.84	4.70	0.000	38	-36	-40	
						0.431	0.075	4.30	4.20	0.000	28	-30	-40	
		0.086	0.097	129	0.005	0.000	0.000	6.01	5.75	0.000	20	-88	-28	
						0.005	0.002	5.48	5.28	0.000	38	-82	-30	
						0.700	0.134	4.07	3.98	0.000	46	-72	-26	
		0.424	0.184	68	0.033	0.002	0.001	5.75	5.52	0.000	-52	-40	-18	
						1.000	0.647	3.34	3.29	0.000	-44	-38	-22	
		0.117	0.097	117	0.007	0.005	0.001	5.51	5.31	0.000	-2	-12	58	
						0.709	0.134	4.06	3.98	0.000	8	-4	52	
		0.393	0.184	71	0.030	0.010	0.003	5.33	5.15	0.000	52	-28	58	
						0.846	0.175	3.93	3.85	0.000	64	-20	42	
		0.857	0.339	34	0.117	0.017	0.004	5.21	5.03	0.000	18	-62	68	
						0.998	0.441	3.53	3.47	0.000	10	-58	66	
		0.210	0.110	95	0.014	0.042	0.010	4.99	4.83	0.000	24	-64	-22	
						0.058	0.013	4.90	4.76	0.000	28	-56	-22	

table shows 3 local maxima more than 8.0mm apart

Height threshold: $T = 3.13$, $p = 0.001$ (1.000)
Extent threshold: $k = 0$ voxels, $p = 1.000$ (1.000)
Expected voxels per cluster, $\langle k \rangle = 14.35$
Expected number of clusters, $\langle c \rangle = 16.6$
FWEp: 4.940, FDRp: 4.534, FWEc: 272, FDRc: 272

Degrees of freedom = [1.0, 198.0]
FWHM = 10.1 10.4 9.6 mm mm mm; 5.0 5.2 4.8 {voxels}
Volume: 1702808 = 212851 voxels = 1574.8 resels
Voxel size: 2.0 2.0 2.0 mm mm mm; (resel = 125.63 voxels)
Page 1

LI based on magnitude

2/5

Statistics: *p-values adjusted for search volume*

set-level		cluster-level				peak-level					mm mm mm		
p	ϵ	$p_{FWL-corr}$	$q_{FDR-corr}$	k_L	p_{uncorr}	$p_{FWL-corr}$	$q_{FDR-corr}$	T	$\{Z\}_\Sigma$	p_{uncorr}			
0.000	39	0.003	0.007	272	0.000	0.000	0.000	8.14	7.55	0.000	30	-36	72
						0.002	0.001	5.72	5.50	0.000	44	-38	64
						0.143	0.027	4.55	4.53	0.000	36	-22	68
		0.592	0.209	54	0.054	0.000	0.000	7.05	6.65	0.000	-36	-20	70
		0.199	0.110	97	0.013	0.000	0.000	7.04	6.64	0.000	-42	-38	62
		0.768	0.311	41	0.088	0.000	0.000	6.41	6.10	0.000	-34	-36	-38
		0.540	0.209	58	0.047	0.000	0.000	6.37	6.07	0.000	-20	-16	76
		0.553	0.209	57	0.048	0.000	0.000	6.24	5.95	0.000	44	-40	-44
						0.073	0.015	4.84	4.70	0.000	38	-36	-40
						0.431	0.075	4.30	4.20	0.000	28	-30	-40

1. Mean maximum T value == mean of top 5% T values

LI based on magnitude

3/5

For each individual participant:

1. Mean maximum T value (T) –

$$T = \text{mean of top 5\% T values } 8.14 - (8.14 * .05) = 7.73$$

2. Take half the mean maximum T value (T/2)

$$T/2 = 7.73/2 = 3.87$$

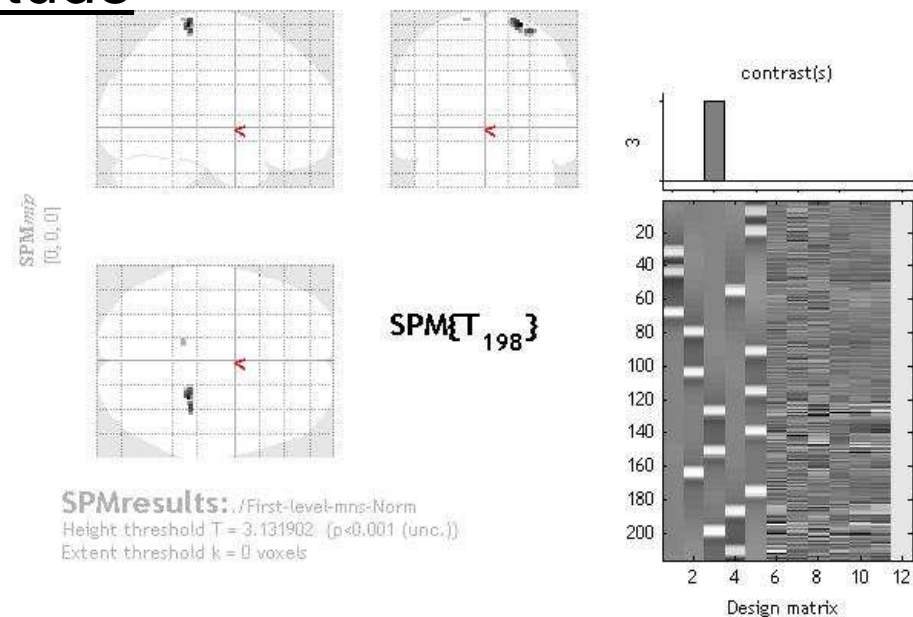
3. In your ROI, sum T values of voxels above $T/2 = (t)$, and calculate LI

$$LI = (t_L -- t_R) / (t_L --+ t_R)$$

see next slide..

LI based on magnitude

4/5



Statistics: *p-values adjusted for search volume*

set-level		cluster-level				peak-level					mm mm mm		
p	c	P _{FWE-corr}	q _{FDR-corr}	k _c	p _{uncorr}	P _{FWE-corr}	q _{FDR-corr}	T	(Z _s)	p _{uncorr}			
1.000	2	0.845	0.339	35	0.112	0.000	0.000	6.37	6.07	0.000	26	-34	72
		1.000	0.772	3	0.653	0.889	0.201	3.88	3.80	0.000	-14	-38	76

3. Inclusive mask of your ROI
OR small volume correct for your ROI–

Here, again using bilateral M1

table shows 3 local maxima more than 8.0mm apart

Height threshold: T = 3.13, p = 0.001 (1,000)	Degrees of freedom = [1.0, 198.0]
Extent threshold: k = 0 voxels, p = 1.000 (1,000)	FWHM = 10.1 10.4 9.6 mm mm mm; 5.0 5.2 4.8 {voxels}
Expected voxels per cluster, <k> = 14.35	Volume: 1702808 = 212851 voxels = 1574.8 resels
Expected number of clusters, <c> = 16.6	Voxel size: 2.0 2.0 2.0 mm mm mm; (resel = 125.63 voxels)
FWEp: 4.940, FDRp: 4.534, FWEc: 272, FDRc: 272	

LI based on magnitude

5/5

Statistics: *p-values adjusted for search volume*

set-level		cluster-level				peak-level					mm mm mm		
p	ϵ	$p_{FWE-corr}$	$q_{FDR-corr}$	k_L	p_{uncorr}	$p_{FWE-corr}$	$q_{FDR-corr}$	T	$(Z_{\frac{\epsilon}{2}})$	p_{uncorr}			
1.000	2	0.845	0.339	35	0.112	0.000	0.000	6.37	6.07	0.000	26	-34	72
		1.000	0.772	3	0.653	0.889	0.201	3.88	3.80	0.000	-14	-38	76

t = sum of t-values that pass $T/2$ in each hemisphere

$$LI = (t_L - t_R) / (t_L + t_R)$$

$$LI = (3.88 - 6.37) / (3.88 + 6.37) = -.24$$

Pretty different from LI calculated based on extent $LI = [-39 \text{ to } -14]$

Last, take mean across par

A. Extent

- Straightforward
- As reproducible

B. Magnitude

- More robust
- Robustness depends on choice of ROI
- Must choose ROI that includes task-related signal,
- E.g. may not be as useful in people with structural l

*slightly different meaning, depending on your question, req

References

1. Cramer, S., Nelles G, Benson RR, Kaplan JD, Parker RA, Kwong KK, Kennedy
A functional MRI study of subjects recovered from hemiparetic stroke. Stroke, 1999. **30**(12): p. 2167-75.
2. Wilke, M. and V.J. Schmithorst, *A combined bootstrap/histogram analysis approach to lateralization index from neuroimaging data.* Neuroimage, 2006. **33**(2): p. 52-61.
3. Wilke, M. and K. Lidzba, *LI-tool: a new toolbox to assess lateralization in functional MRI.* J Neurosci Methods, 2007. **163**(1): p. 128-36.
4. Fernandez, G., et al., *Language mapping in less than 15 minutes: real-time functional MRI investigation.* Neuroimage, 2001. **14**(3): p. 585-94.
5. Jansen, A., et al., *The assessment of hemispheric lateralization in functional MRI.* Neuroimage, 2006. **33**(1): p. 204-17.