

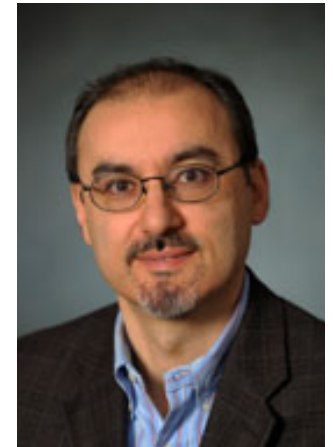
# Harmonizing sMRI Data via Robust Preprocessing

Aristeidis Sotiras

Postdoctoral Research Fellow

Section of Biomedical Image Analysis

# SBIA



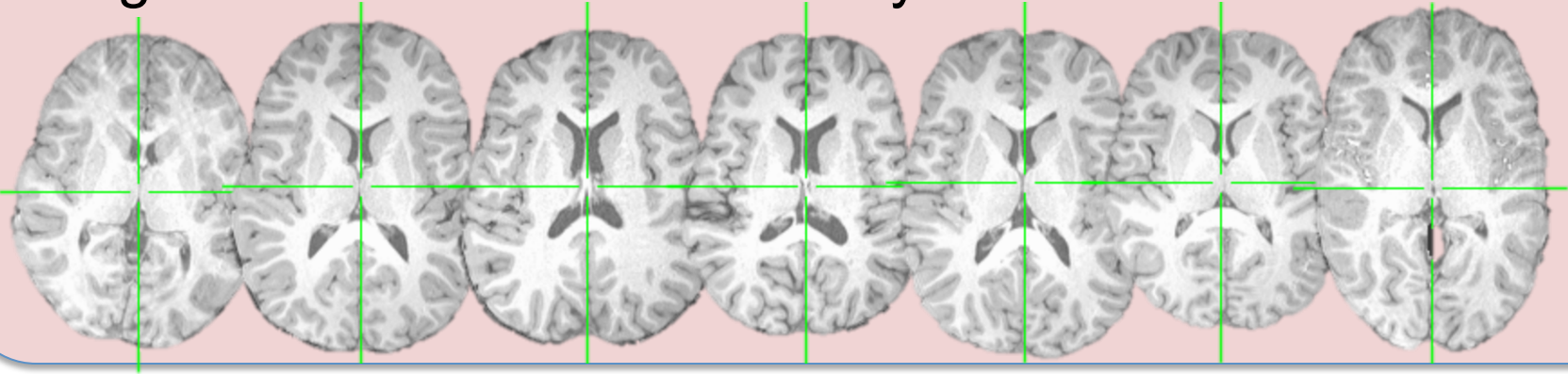
# SBIA – Neuroimaging Projects

Study	Multi-site	Scanner	Participants
BLSA	No	1.5T/3T	160
SCZ	Yes	1.5T/3T	1081
ADNI	Yes	1.5T	822
PNC	No	3T	1,445
ACCORD	Yes	1.5T	729 (BL) – 511 (FUP)
CARDIA	Yes	3T	~600
NiCK	No	3T	180
Sprint	Yes	3T	640

- “*Big Data*” projects:
- high sample size
  - multi-site
  - different imaging protocols

# Challenges

- Significant anatomical variability



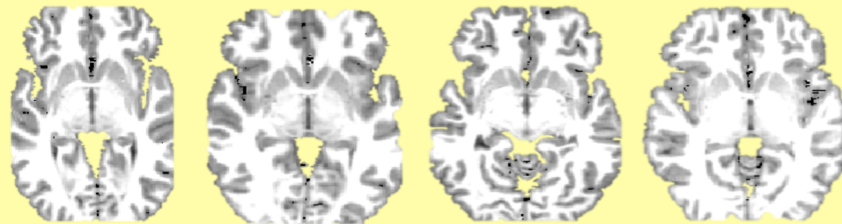
- Significant differences in intensity characteristics



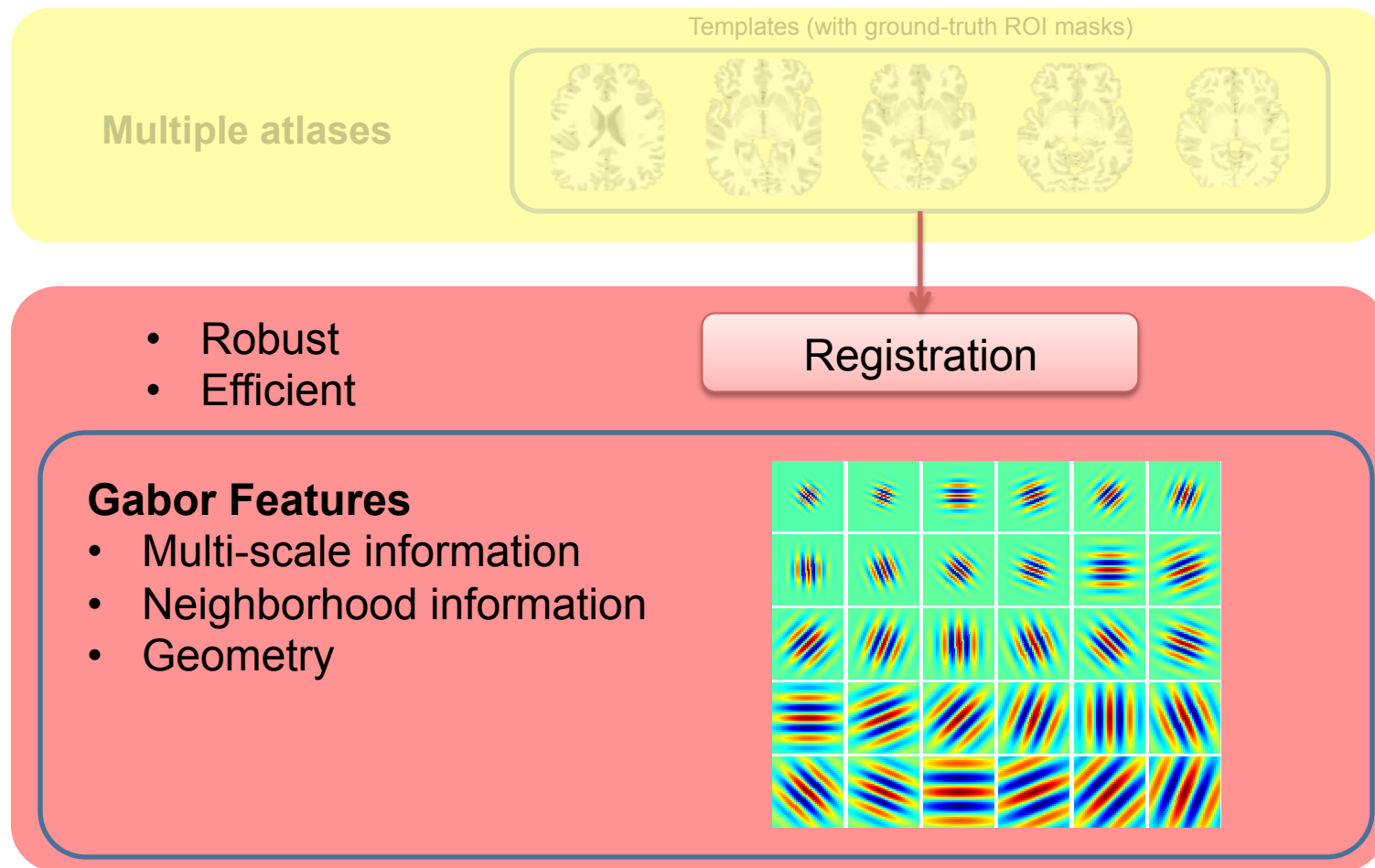
# Robust Preprocessing based on Multi-Atlas Framework

- Multiple atlases** capturing
- Anatomical variability
  - Intensity variability

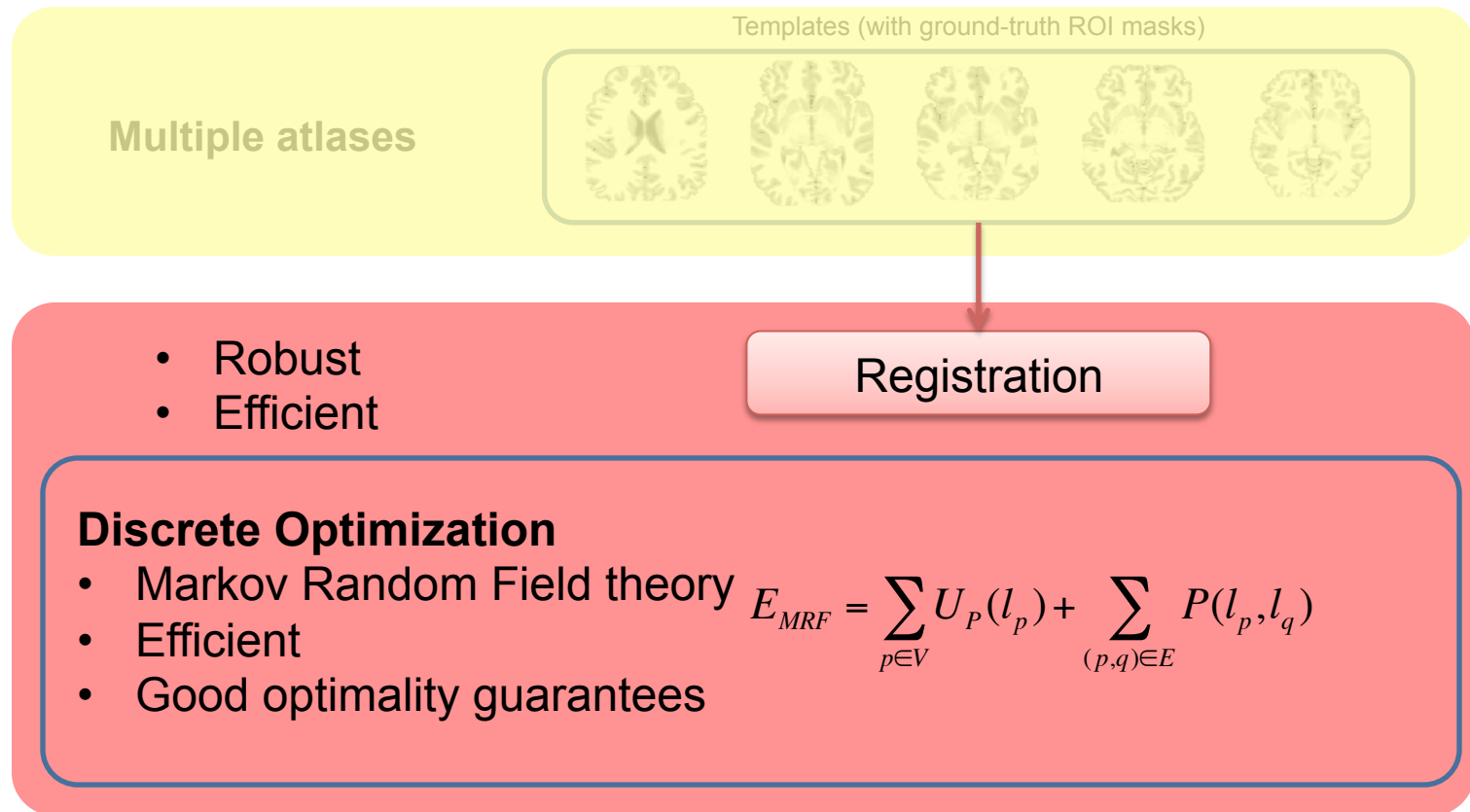
Templates (with ground-truth ROI masks)



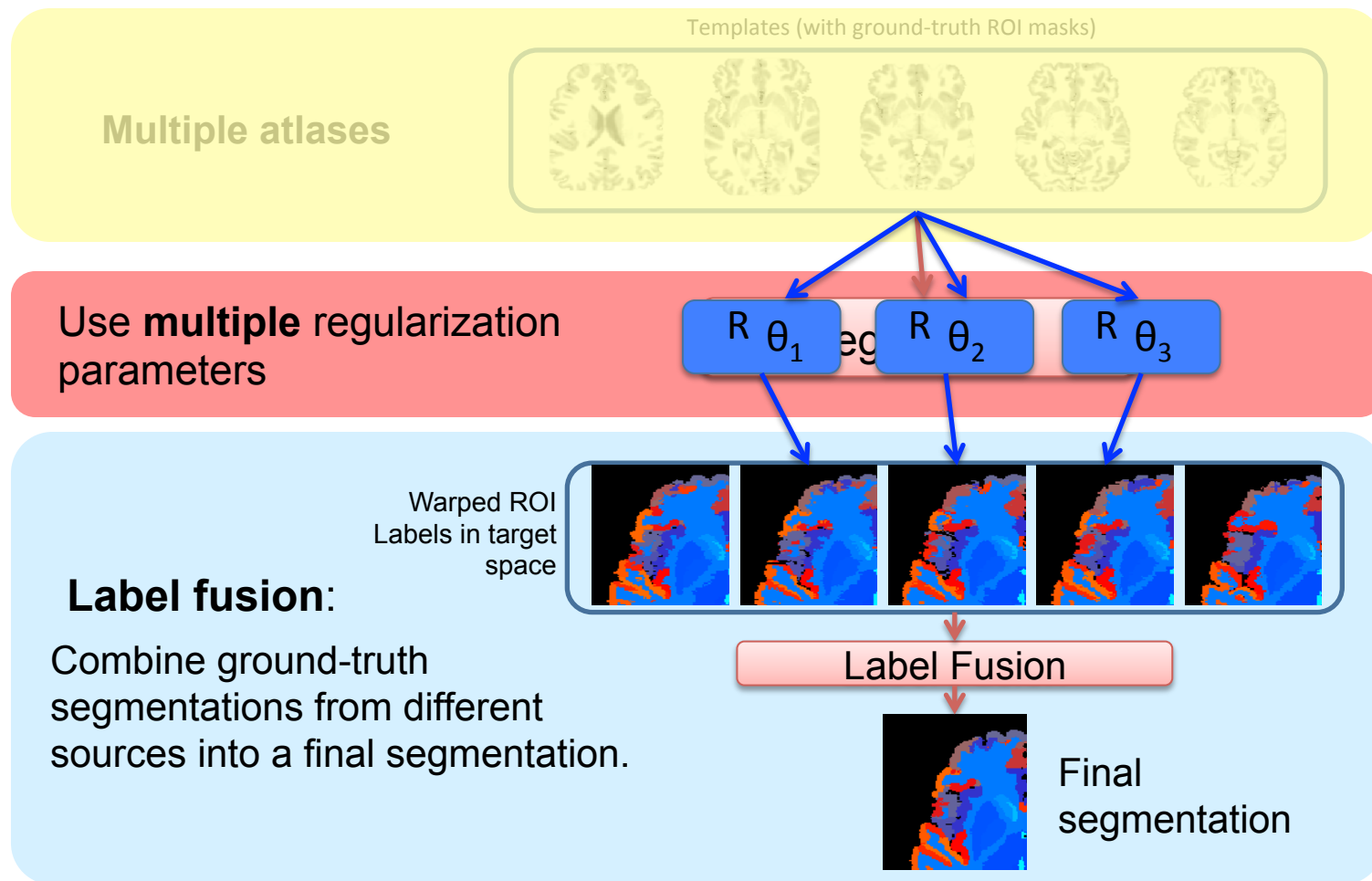
# Robust Preprocessing based on Multi-Atlas Framework



# Robust Preprocessing based on Multi-Atlas Framework

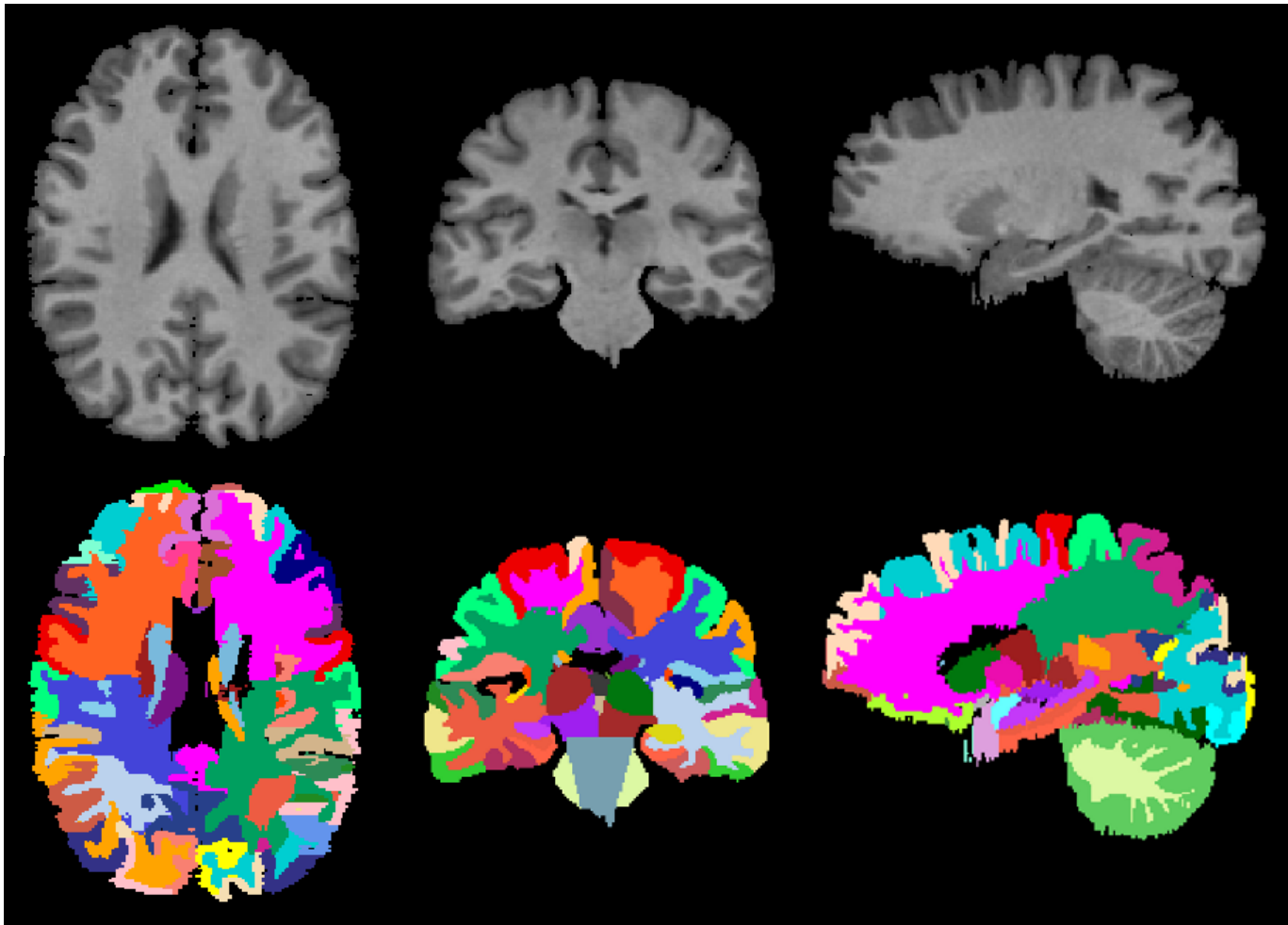


# Robust Preprocessing based on Multi-Atlas Framework





# Hierarchical Multi-Atlas Representation



Doshi, Jimit, et al. "Multi-atlas skull-stripping."  
*Academic radiology* 20.12 (2013): 1566-1576.

# Hierarchical Multi-Atlas Representation

TOTALBRAIN	TISSUE_SEG	SUBGROUP_0	SUBGROUP_1	SUBGROUP_2	ROI_NAME
	GM	FRONTAL	FRONTAL_GM	FRONTAL_INFERIOR_GM	Left AOrG anterior orbital gyrus
	GM				Left LOrG lateral orbital gyrus
	GM				Right MOrG medial orbital gyrus
	GM				Right POrG posterior orbital gyrus
	GM			FRONTAL_INSULAR_GM	Left AIns anterior insula
	GM				Left PIns posterior insula
	GM				Right AIns anterior insula
	GM				Right PIns posterior insula
	GM			FRONTAL_LATERAL_GM	Right OpIFG opercular part of the inferior frontal gyrus
	GM				Right OrIFG orbital part of the inferior frontal gyrus
	GM				Right PrG precentral gyrus
	GM				Right SFG superior frontal gyrus
	GM			FRONTAL_MEDIAL_GM	Right TriFG triangular part of the inferior frontal gyrus
	GM				Left GRe gyrus rectus
	GM				Left MFC medial frontal cortex
	GM				Left MPrG precentral gyrus medial segment
	GM				Right MPrG precentral gyrus medial segment
	GM				Right MSFG superior frontal gyrus medial segment
	GM			FRONTAL_OPERCULAR_GM	Right SCA subcallosal area
	GM				Right SMC supplementary motor cortex
	GM				Left CO central operculum
	GM				Left FO frontal operculum
	GM				Left PO parietal operculum
	GM				Right CO central operculum
	GM	Right FO frontal operculum			
	GM	Right PO parietal operculum			
	WM	FRONTAL_WM	frontal lobe wM left		
	WM		frontal lobe wM right		
	GM	PARIETAL	PARIETAL_LATERAL_GM	Left AnG angular gyrus	
	GM			Left PoG postcentral gyrus	
	GM			Right PoG postcentral gyrus	
	GM			Right SMG supramarginal gyrus	
	GM			Right SPL superior parietal lobule	
	GM			Left MPoG postcentral gyrus medial segment	
	GM		PARIETAL_MEDIAL_GM	Left PCu precuneus	
	GM			Right MPoG postcentral gyrus medial segment	
	GM			Right PCu precuneus	
	WM			parietal lobe wM left	
	WM	PARIETAL_WM	parietal lobe wM right		

TOTALBRAIN

# Harmonizing BLSA\*

\*BLSA: Baltimore Longitudinal Study of Aging  
America's longest-running scientific study of human aging  
Longitudinal scans of hundreds of subjects

Main **challenge**:

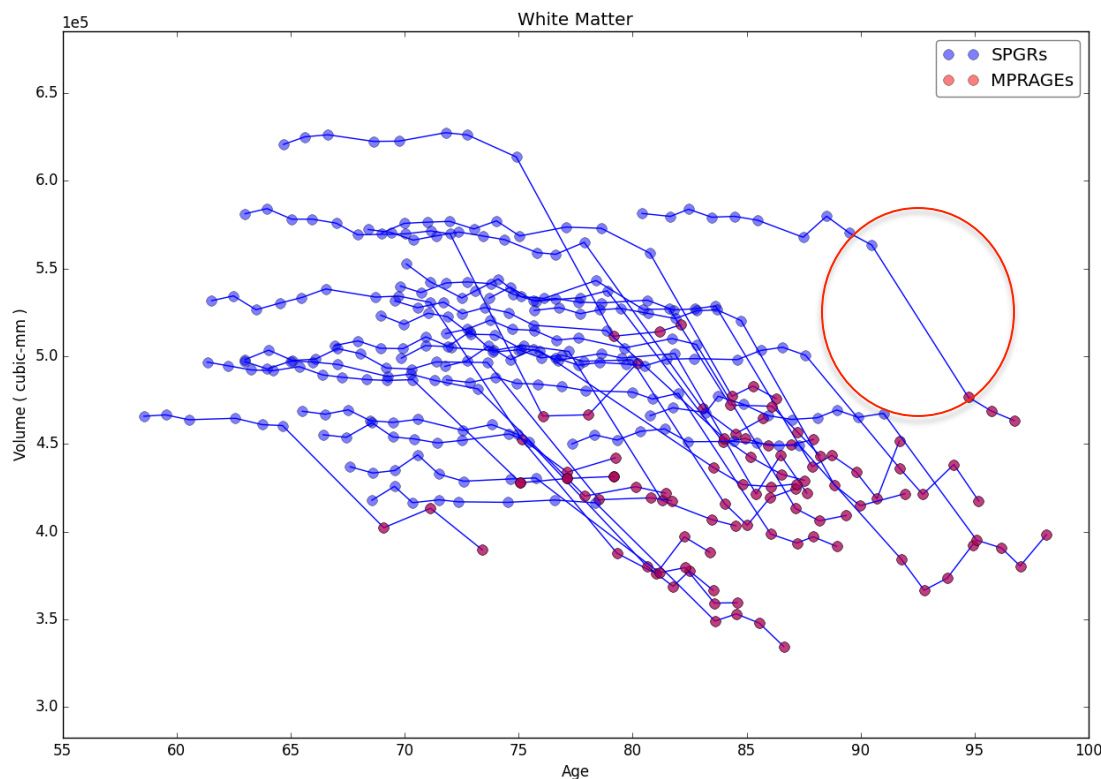
Scanner change from 1.5T SPGR to 3T MPRAGE

Scanner	1.5T	3T
Participants	160	561
Gender: males (females)	92 (68)	252 (309)
Age: mean $\pm$ std (range)	73.79 $\pm$ 8.17 (45.64)	71.87 $\pm$ 13.00 (69.33)
No of scans	1089	862

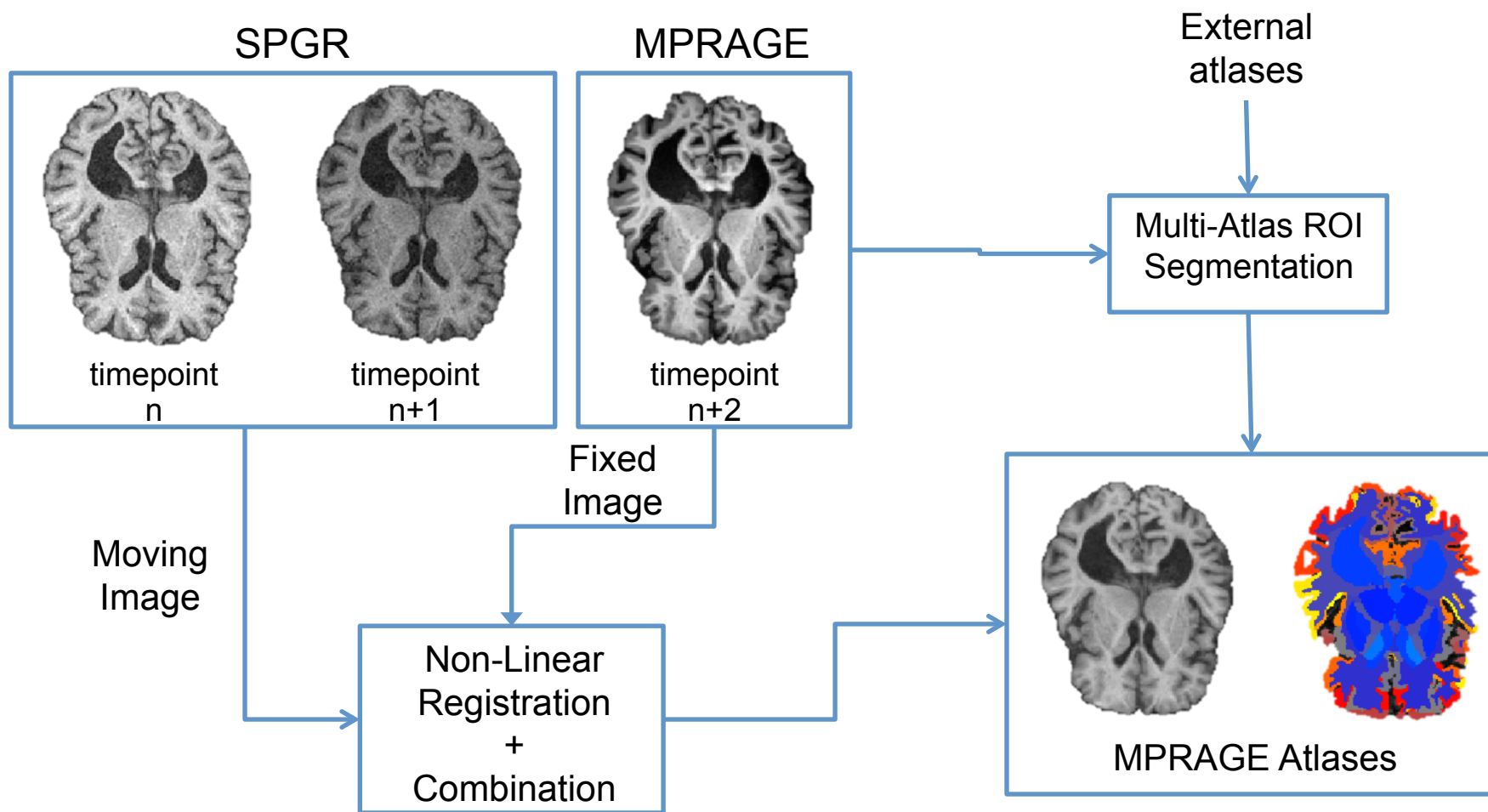
# Harmonizing BLSA

Image contrast differences between SPGR and MPRAGE:

- Lead to **under-segmentation**
- **Solution**: adaptive atlas construction

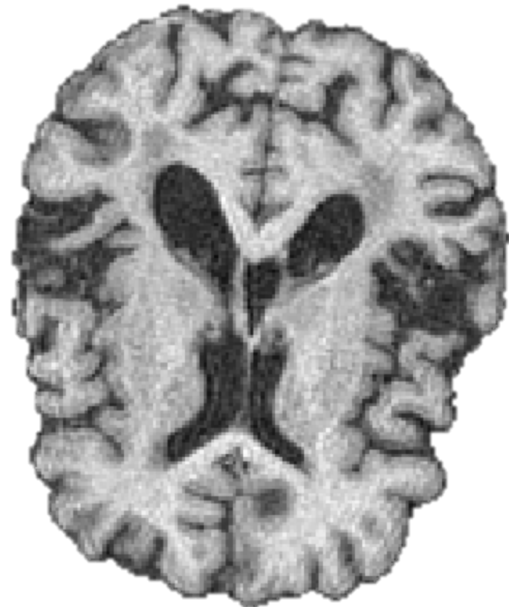


# Adaptive Atlas Construction

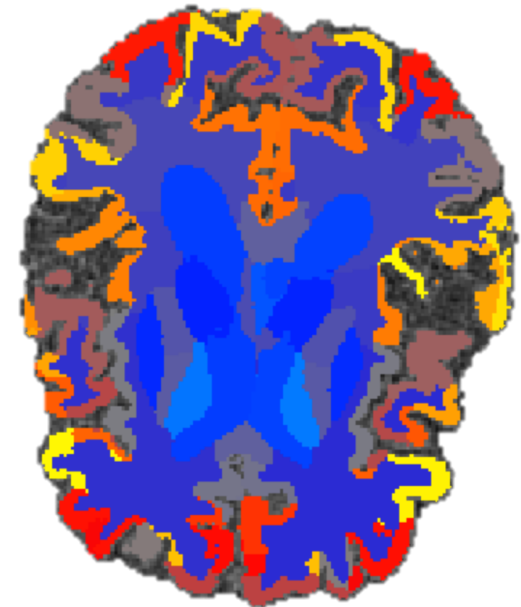
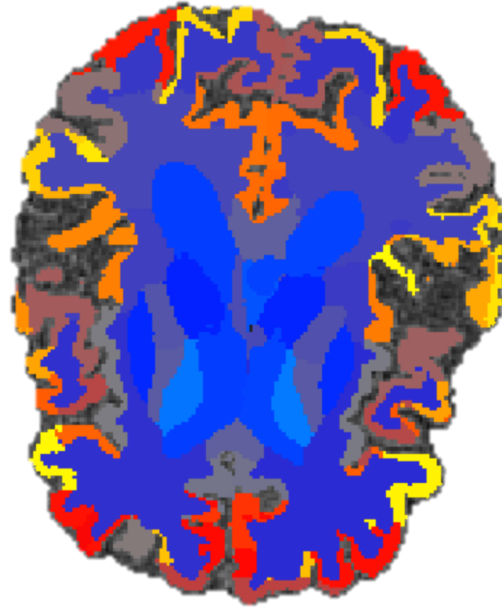


# Segmentation Results

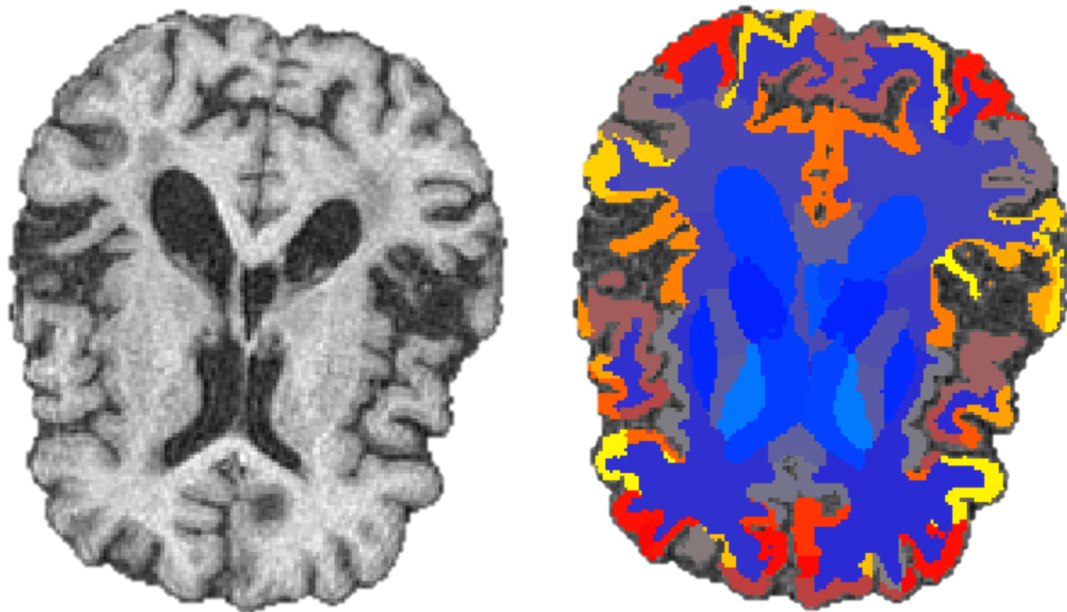
Previous  
Segmentation



Adaptive  
Segmentation



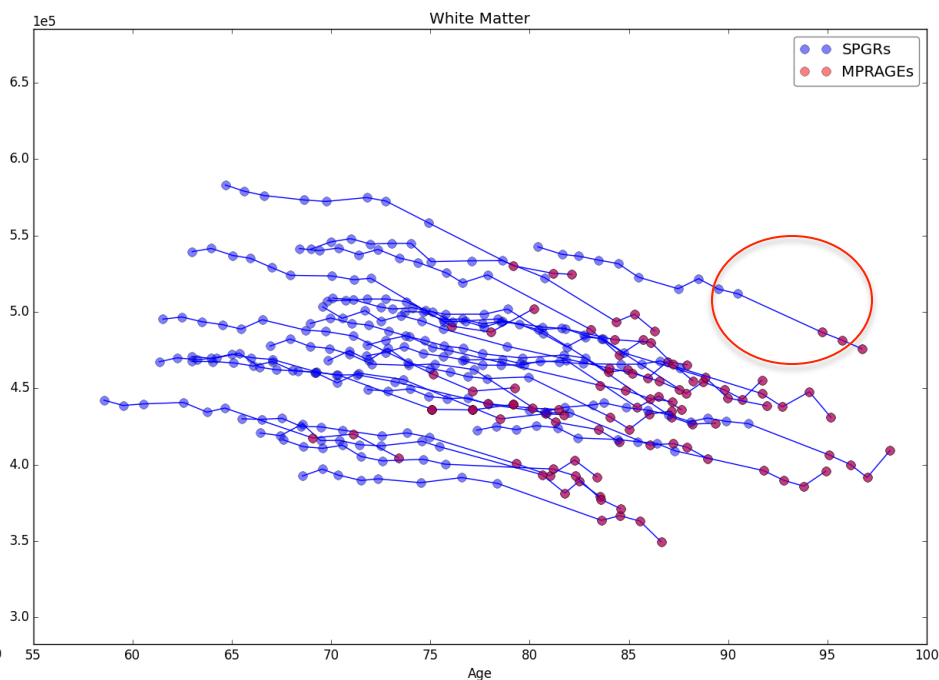
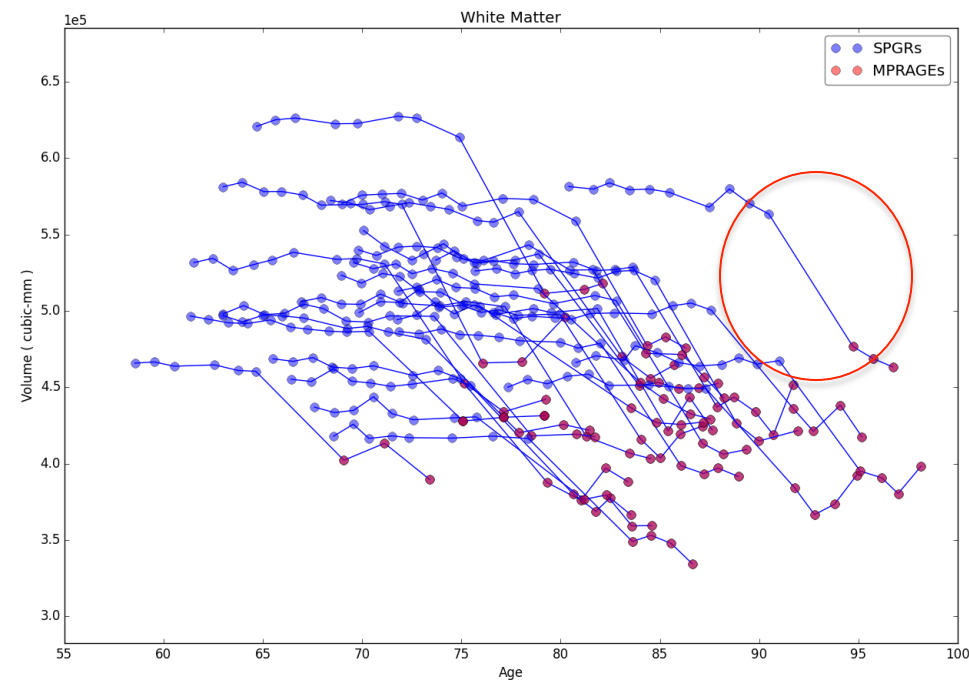
# Segmentation Results



# Longitudinal WM Trajectories

Initial

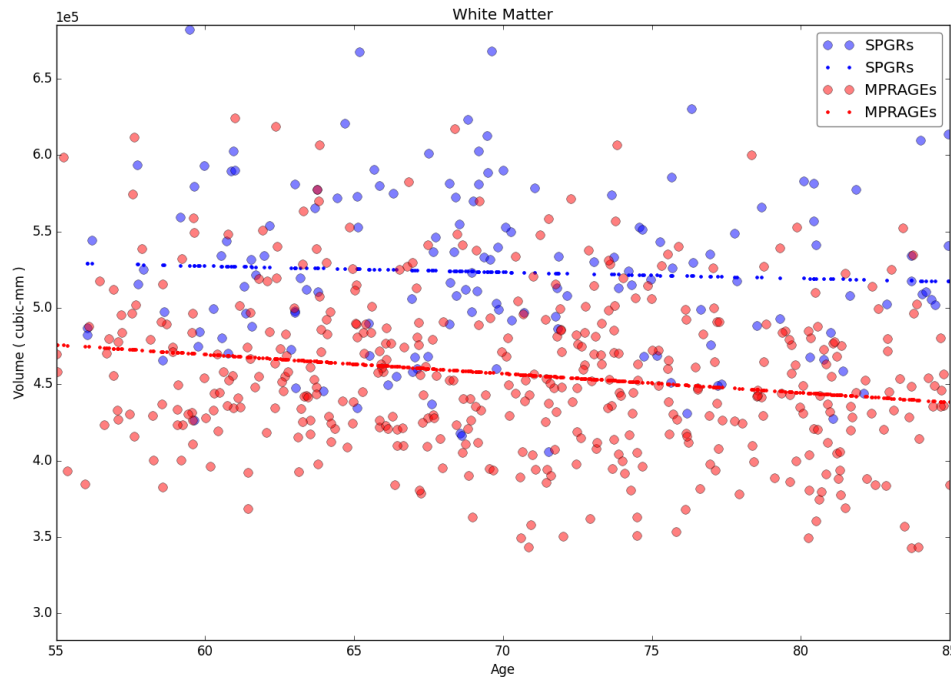
Adaptive



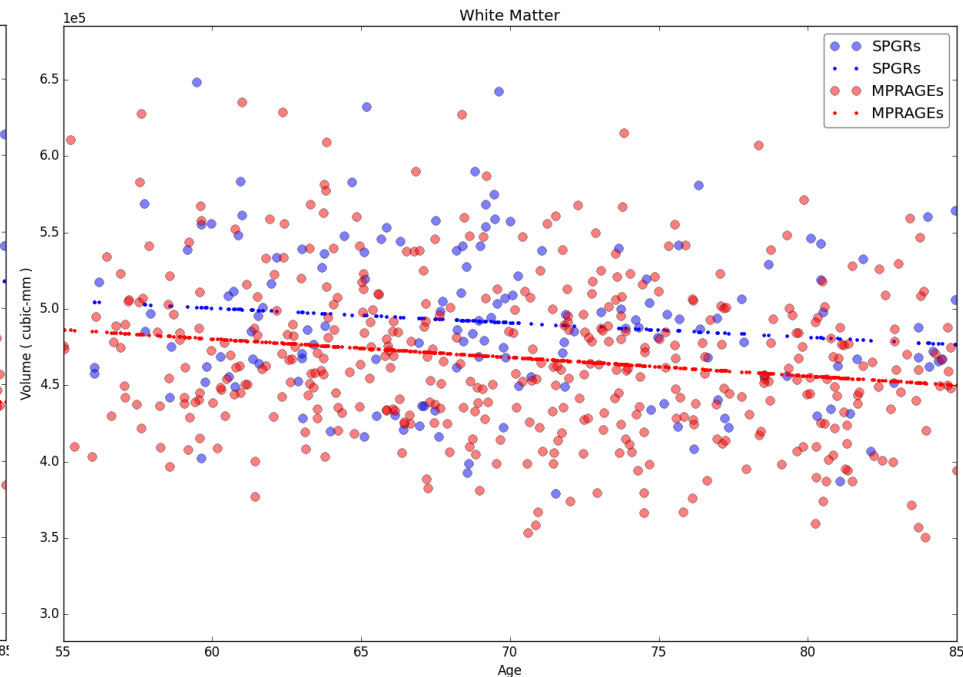


# Cross-Sectional WM Trends

Initial



Adaptive

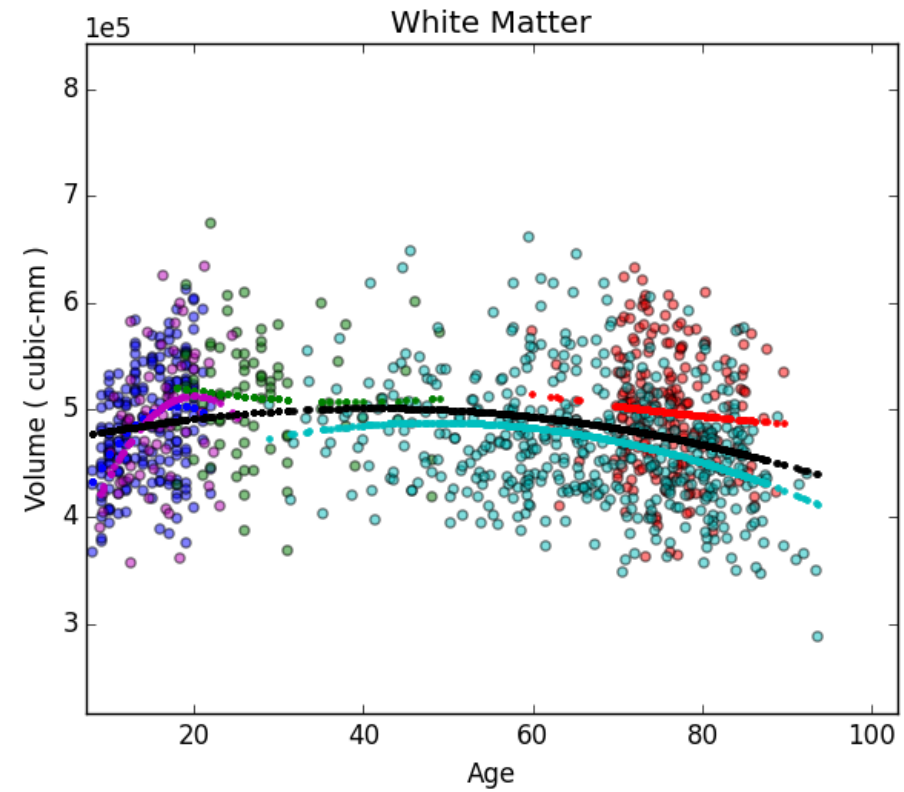
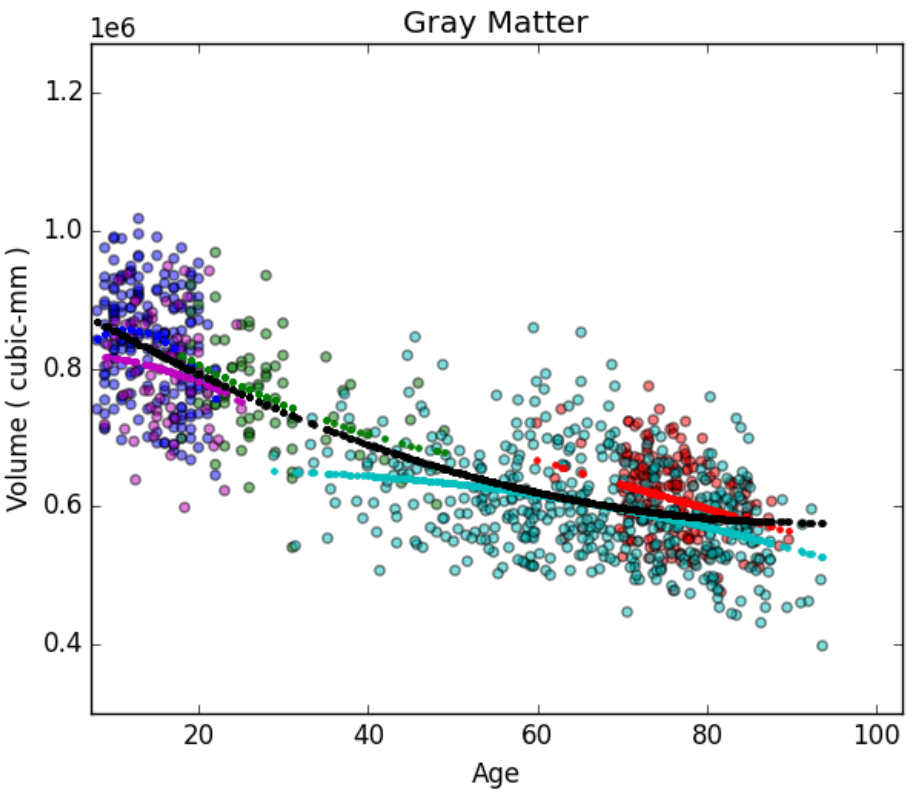


# Normative Brain Dataset

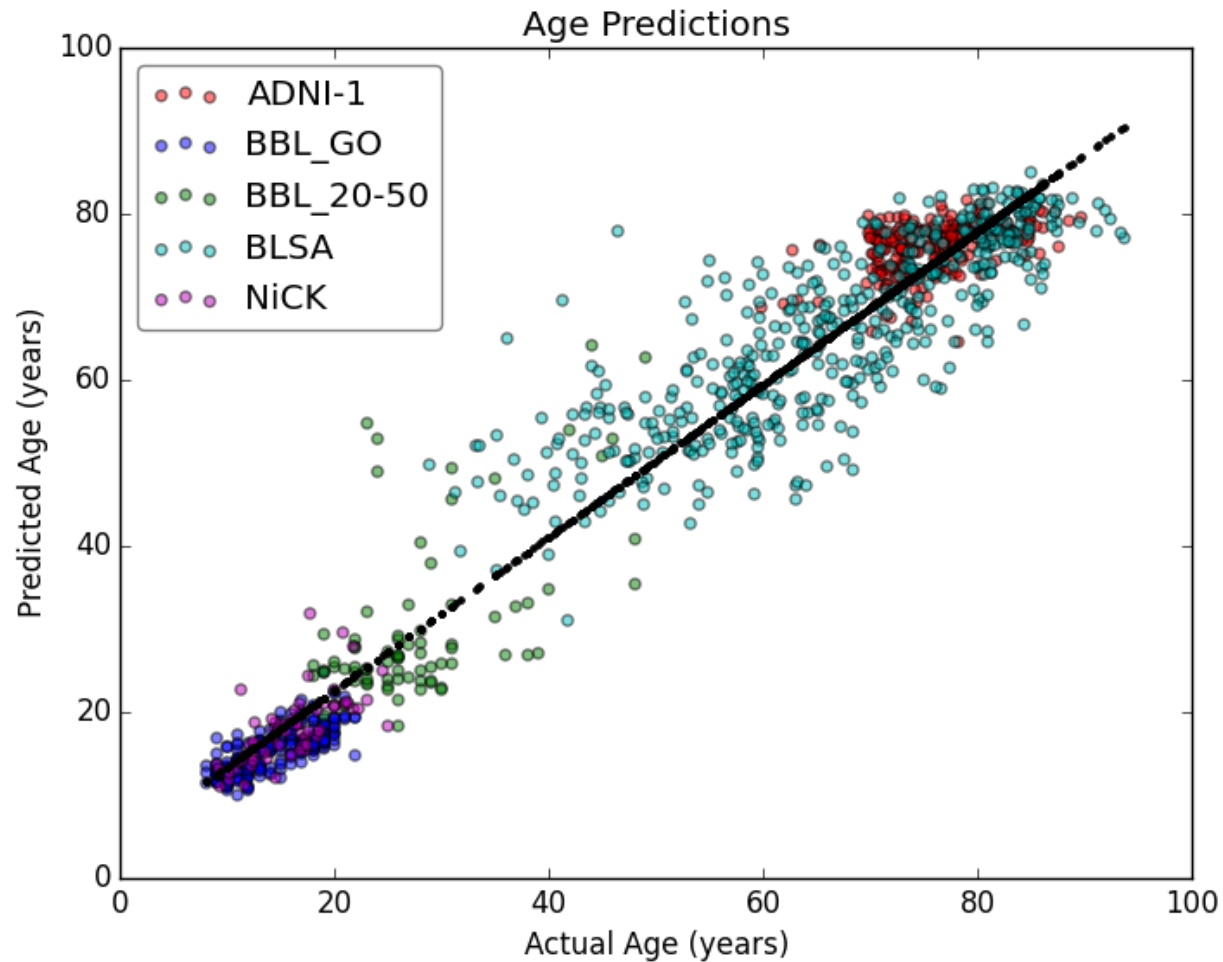
**Main objective:** Combining control subjects from different studies and extracting robust imaging features for the analysis of structural brain age-related change across life span.

Study	Scanner	Subjects	Gender	Age (Years)
			Male/Female	Mean $\pm$ SD (range)
BLSA	1.5T	468	220/248	66.40 $\pm$ 13.91 (65)
BBL_GO	3T	201	101/100	14.51 $\pm$ 3.79 (14)
BBL_20-50	1.5T	79	41/38	28.22 $\pm$ 7.52 (31)
ADNI	1.5T	215	109/106	75.83 $\pm$ 5.03 (30)
NiCK	3T	66	35/31	15.89 $\pm$ 3.90 (16)

# ROI Volumes across Life Span



# Age Prediction Across Life Span

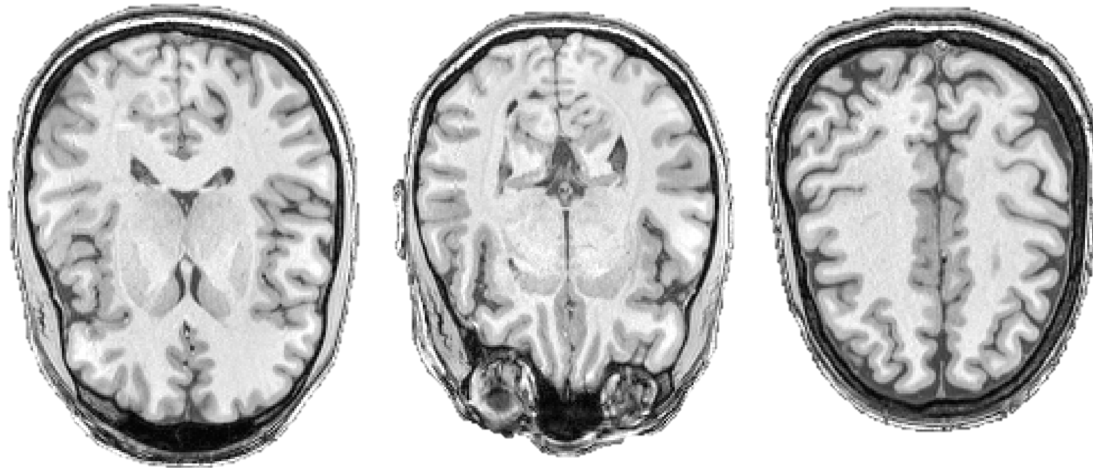


# Harmonizing SCZ Study

Data Site	Scanner	Diagnosis	Subjects	Gender	Age (Years)
				Male/Female	Mean $\pm$ SD (range)
<b>Penn</b>	3T	SCZ	138	86/52	35.38 $\pm$ 11.34 (13-60)
		NC	132	63/69	31.80 $\pm$ 12.89 (12-65)
<b>China</b>	3T	SCZ	144	73/71	30.35 $\pm$ 9.52 (14-59)
		NC	169	79/90	31.63 $\pm$ 10.54 (17-57)
<b>Munich</b>	1.5T	SCZ	165	123/42	31.35 $\pm$ 9.66 (18-65)
		NC	177	123/54	31.48 $\pm$ 9.16 (18-61)
<b>Brazil</b>	1.5T	SCZ	62	45/17	27.74 $\pm$ 8.00 (18-50)
		NC	94	53/41	30.21 $\pm$ 8.40 (18-50)
<b>TOTAL</b>		SCZ	509	327/182	31.73 $\pm$ 10.18 (13-65)
		NC	572	318/254	31.39 $\pm$ 10.41 (12-65)

# Challenges: Contrast Difference

Penn

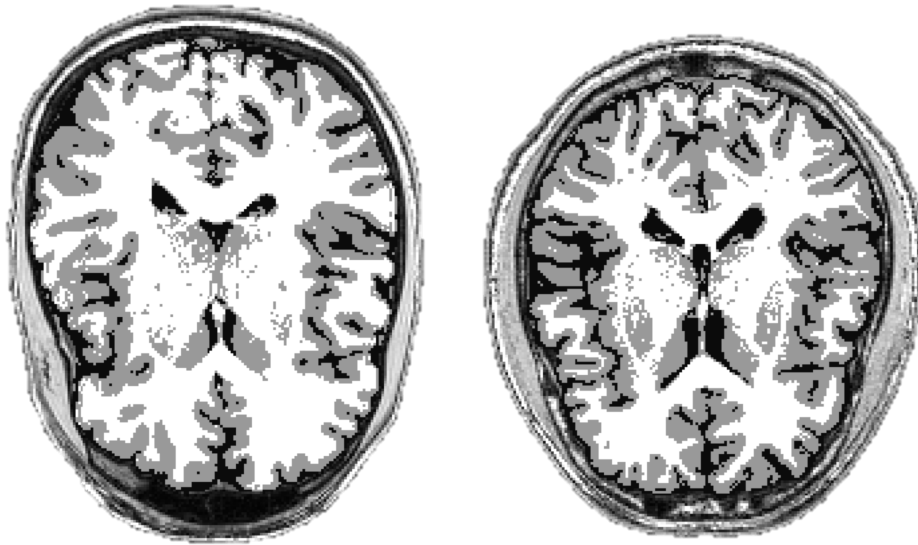


China

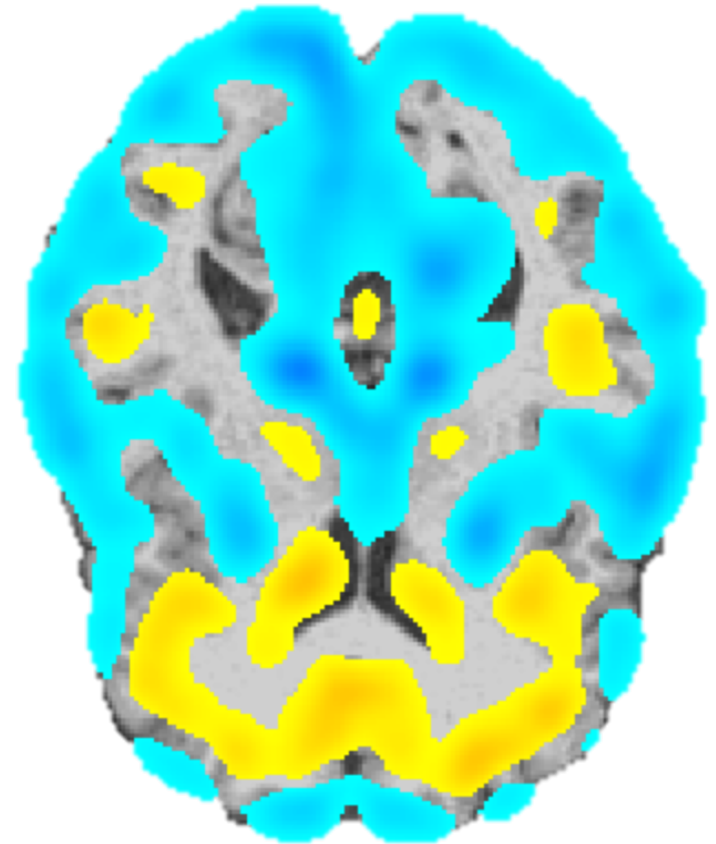


# Dataset Differences

GM-WM-CSF Segmented Images



VBM comparison of dataset control GM

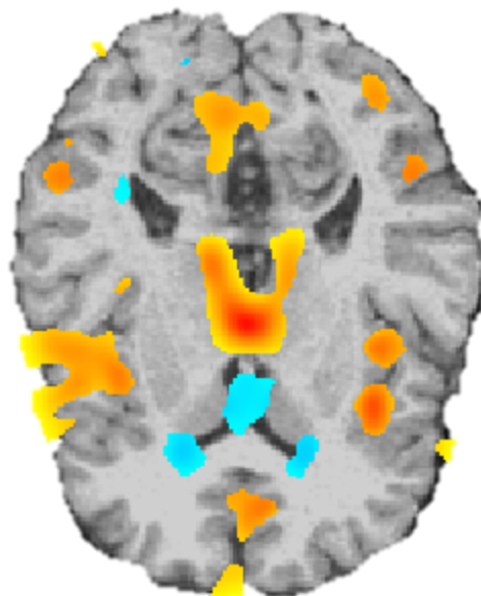
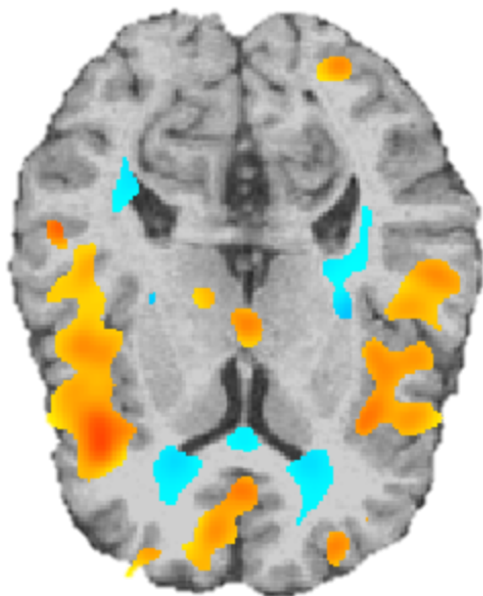


Blue: Penn less GM  
Yellow: Penn more GM

# Harmonizing by CN Means

**VBM Comparison of NC vs SCZ  
within each dataset**

**VBM Comparison of NC vs SCZ for  
Pooled Datasets**

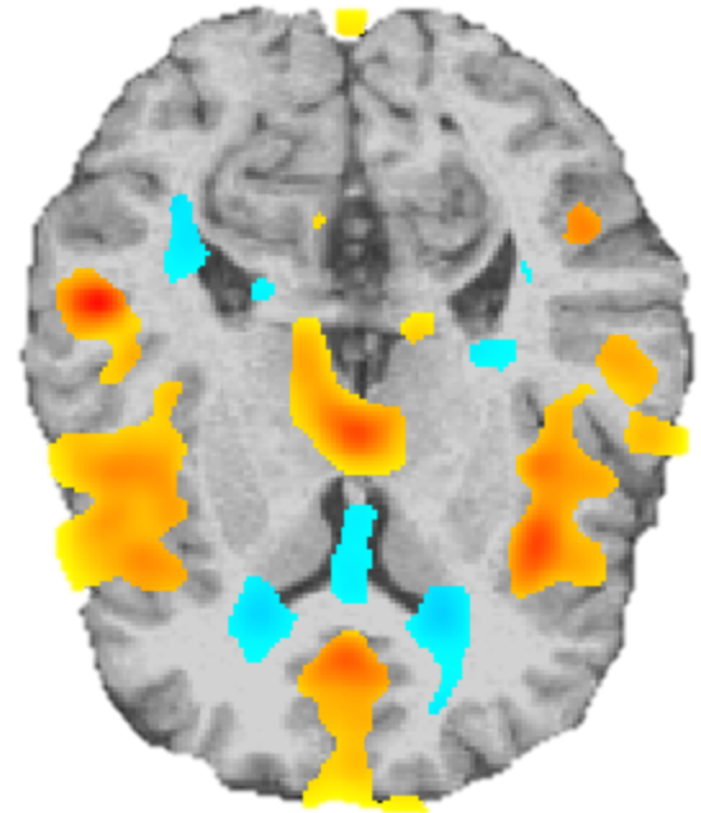


Penn

China

**Key**

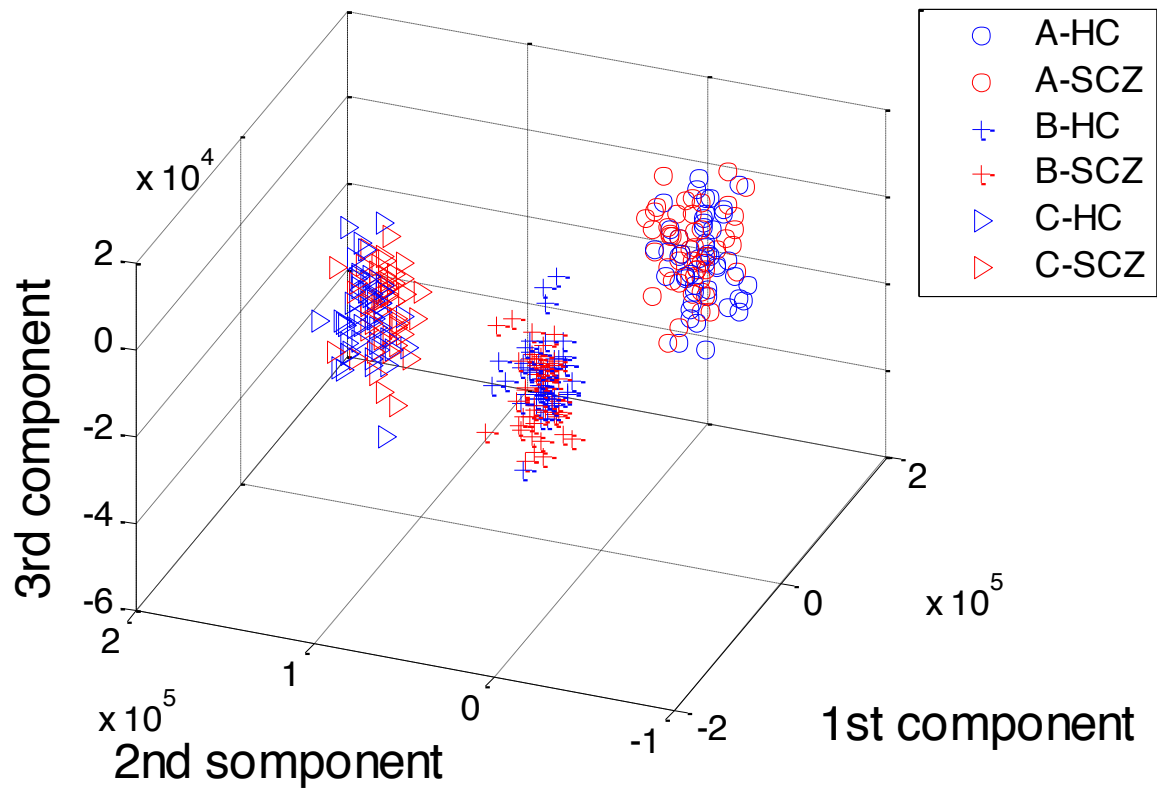
Blue: Control GM < SCZ GM  
Yellow: Control GM > SCZ GM





# Classification Accuracy

Demographic information



Classification accuracy

Site	SVM	GLM+SVM
A	72	76
B	67	74
C	68	84

a: Pearson Chi-square test. b: Two-sample t-test.

# Conclusions

- Important challenges associated with harmonizing shared resources
  - Anatomical variability
  - Different intensity properties
- Robust preprocessing removes important confounding variation
- Simple statistical models help
- Can we do more ?
  - Machine learning approaches based on multi-task learning
  - Machine learning techniques that model explicitly the confounding factors

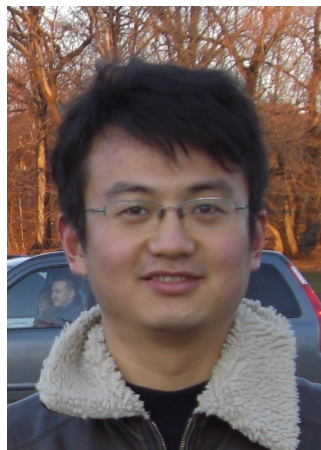
# Credits



C. Davatzikos



G. Erus



T. Zhang



J. Doshi



M. Rozycki



Q. Ma

