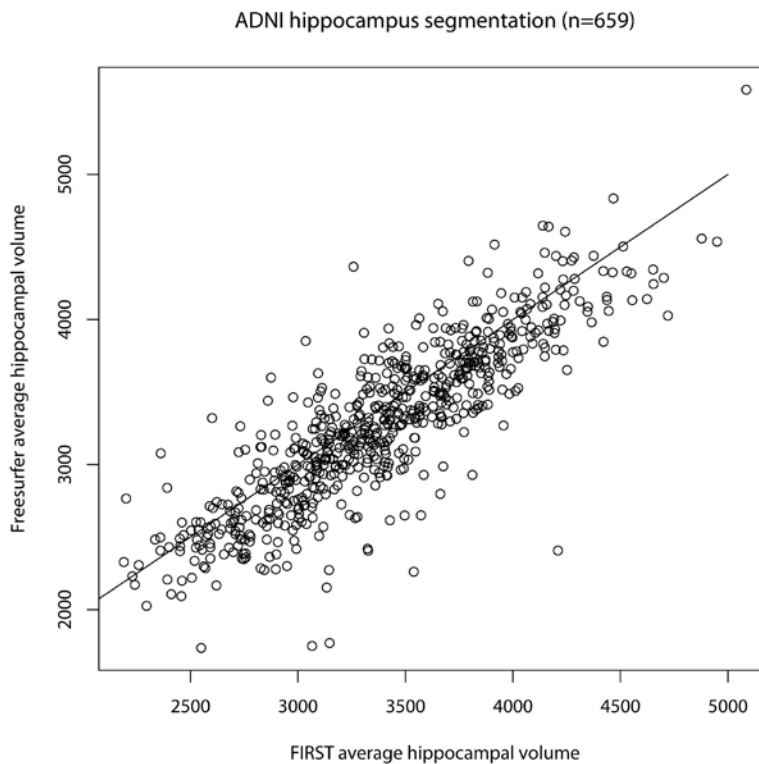


## Freesurfer vs. FIRST

### *ADNI Sample:*

We conducted a direct comparison of Freesurfer and FIRST segmentation of the hippocampus in the ADNI sample. Freesurfer volumes were downloaded from the LONI IDA and FIRST segmentation was completed according to the protocol on the ENIGMA website (<http://enigma.loni.ucla.edu/protocols/>). Upon removing segmentations which contained visually incorrect segmentations, the overlapping number of subjects was n=659 which included healthy elderly subjects, mild cognitively impaired, and Alzheimer's disease patients. The average bilateral hippocampal volume yields Pearson's correlation of 0.86 between segmentation criteria.

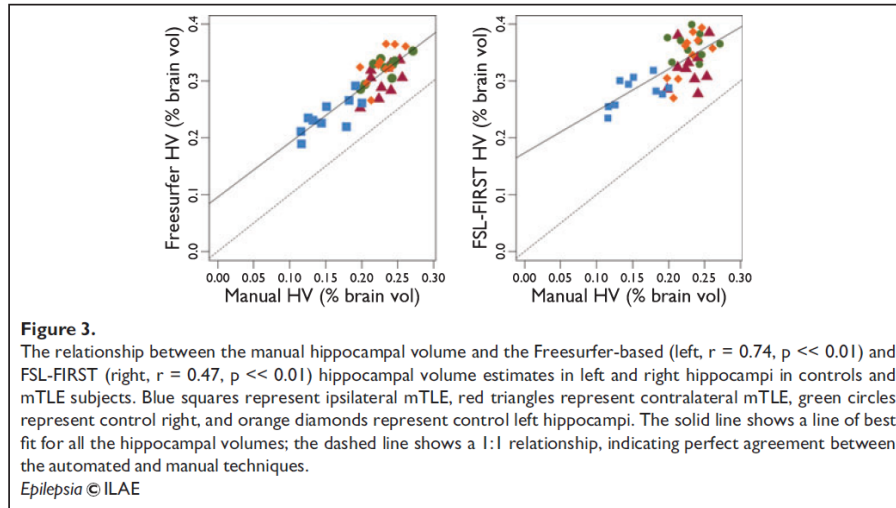


### *Work comparing methods done by others:*

In other work, Freesurfer has a higher correlation to manual segmentations than FIRST. The overlap between Freesurfer/FIRST is  $0.72 \pm 0.037$  in controls and  $0.67 \pm 0.042$  in temporal lobe epilepsy patients (Pardoe et al., 2009).

<b>Table 2. Overlap (Dice coefficient) between different hippocampal segmentation methods in controls and left mTLE hippocampi</b>		
	Controls (mean $\pm$ SD)	mTLE (mean $\pm$ SD)
Manual/Freesurfer	$0.73 \pm 0.028$	$0.66 \pm 0.042$
Manual/FSL-FIRST	$0.71 \pm 0.046$	$0.62 \pm 0.057$
Freesurfer/FSL-FIRST	$0.72 \pm 0.037$	$0.67 \pm 0.042$

The overlap is reduced in affected mTLE hippocampi, indicating less agreement between manual and automated methods ( $p < 0.05$  for each control/mTLE comparison).



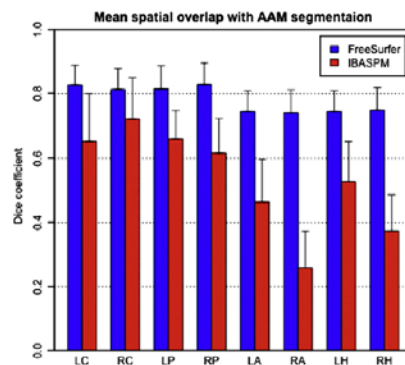
Freesurfer is more accurate than FIRST according to (Morey et al., 2009a). There was a small controversy about this because the authors did not report enough demographic information (Hasan and Pedraza, 2009), however it has not changed the results (Morey et al., 2009b).

**Table 1**  
Comparison of automated measures to manual tracing

Automated measure	Average volume $\pm$ SD	% Volume overlap		% Volume difference		Correlation with manual segmentation
		Left	Right	Left	Right	
<i>Hippocampus</i>						
FreeSurfer	4190 $\pm$ 526.7	82% $\pm$ 1.5	82% $\pm$ 2.8	4% $\pm$ 2.1	5% $\pm$ 1.7	$r = 0.82$ , y-intercept = 496
FSL-FIRST (threshold = 2)	4193 $\pm$ 634.9	79% $\pm$ 3.6	80% $\pm$ 2.9	4% $\pm$ 2.4	4% $\pm$ 2.3	$r = 0.66$ , y-intercept = 1502
FSL-FIRST (threshold = 3)	4843 $\pm$ 743.4	78% $\pm$ 3.8	79% $\pm$ 2.8	6% $\pm$ 3.4	7% $\pm$ 2.3	$r = 0.66$ , y-intercept = 1480
FSL-FIRST (version 4.1)	4404 $\pm$ 730.1	77% $\pm$ 5.9	80% $\pm$ 2.4	5% $\pm$ 3.2	5% $\pm$ 2.8	$r = 0.66$ , y-intercept = 1480

One study shows that Freesurfer is less accurate than HAMMER (Akhondi-Asl et al., 2010), though it look like one of the segmentations failed completely in several cases in this study and that was not corrected.

Freesurfer is more accurate than Individual Brain Atlases using Statistical Parametric Mapping (IBASPM) (Dewey et al., 2010).



**Fig. 1.** FreeSurfer achieved significantly higher spatial overlap with AAM segmentation in every structure. Error bars indicate one standard deviation.

Another study shows that FreeSurfer is fairly accurate but overestimates the volumes (Sanchez-Benavides et al., 2010).

**Table 3**  
Comparison of manual and automated volumetric measures of hippocampus.

Manual vs. FreeSurfer	Global	Controls	MCI	AD
<b>Left hippocampus</b>				
% Overlap	79% ( $\pm 7$ )	81% ( $\pm 5$ )	79% ( $\pm 7$ )	75% ( $\pm 9$ )
% Difference	10% ( $\pm 7$ )	9% ( $\pm 8$ )	10% ( $\pm 7$ )	11% ( $\pm 12$ )
Pearson correlation	0.85	0.74	0.82	0.84
<b>Right hippocampus</b>				
% Overlap	77% ( $\pm 6$ )	80% ( $\pm 4$ )	76% ( $\pm 6$ )	74% ( $\pm 7$ )
% Difference	11% ( $\pm 8$ )	9% ( $\pm 7$ )	9% ( $\pm 8$ )	13% ( $\pm 8$ )
Pearson correlation	0.84	0.74	0.79	0.71

Overlap and difference percentages, and Pearson correlation coefficient.  
MCI: Mild Cognitive Impairment; AD: Alzheimer Disease.

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