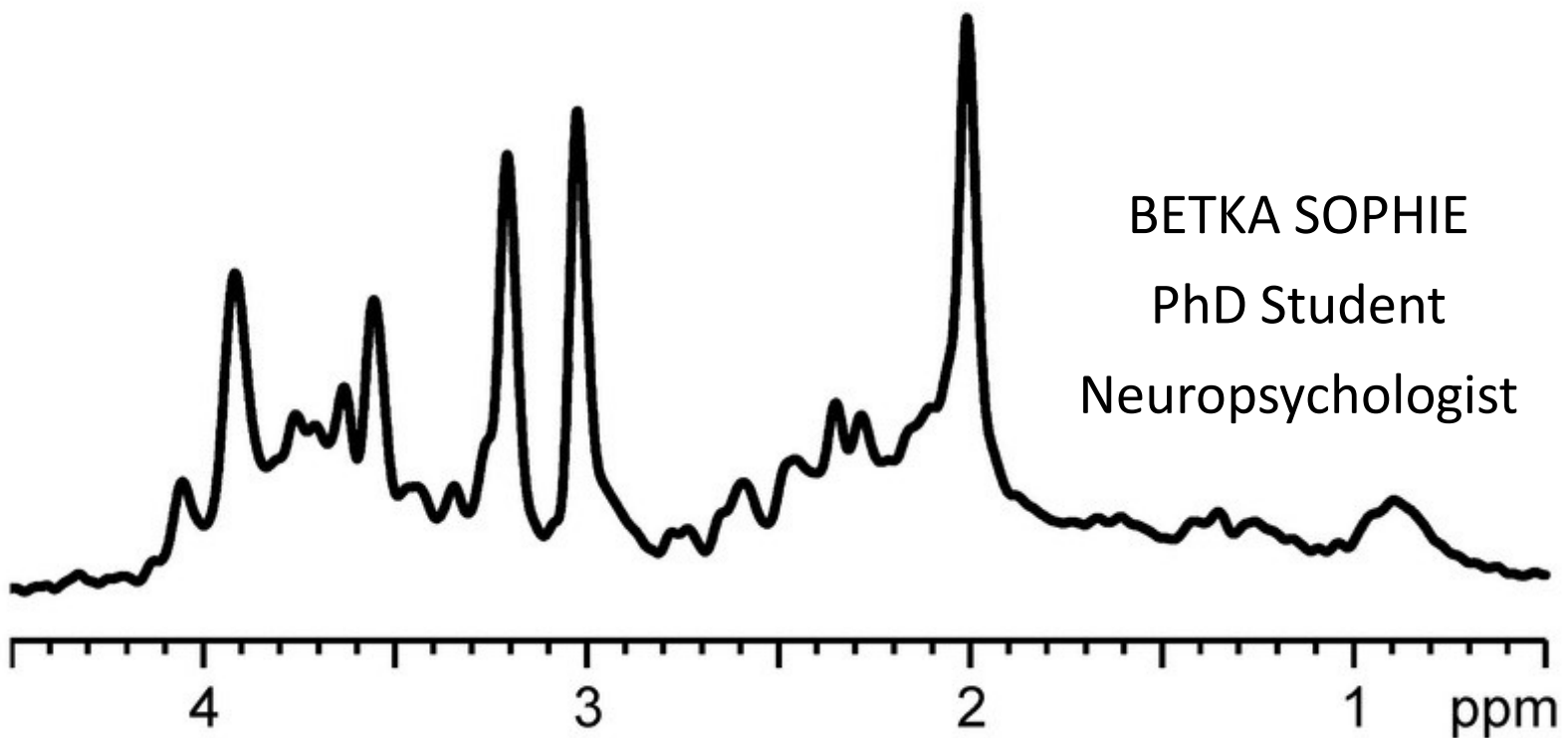


# Bodily sensation processing in social drinking: *a combined MRS, VBM & SBM study*

*Society for the Study of Addiction*

*10<sup>th</sup> November 2017*



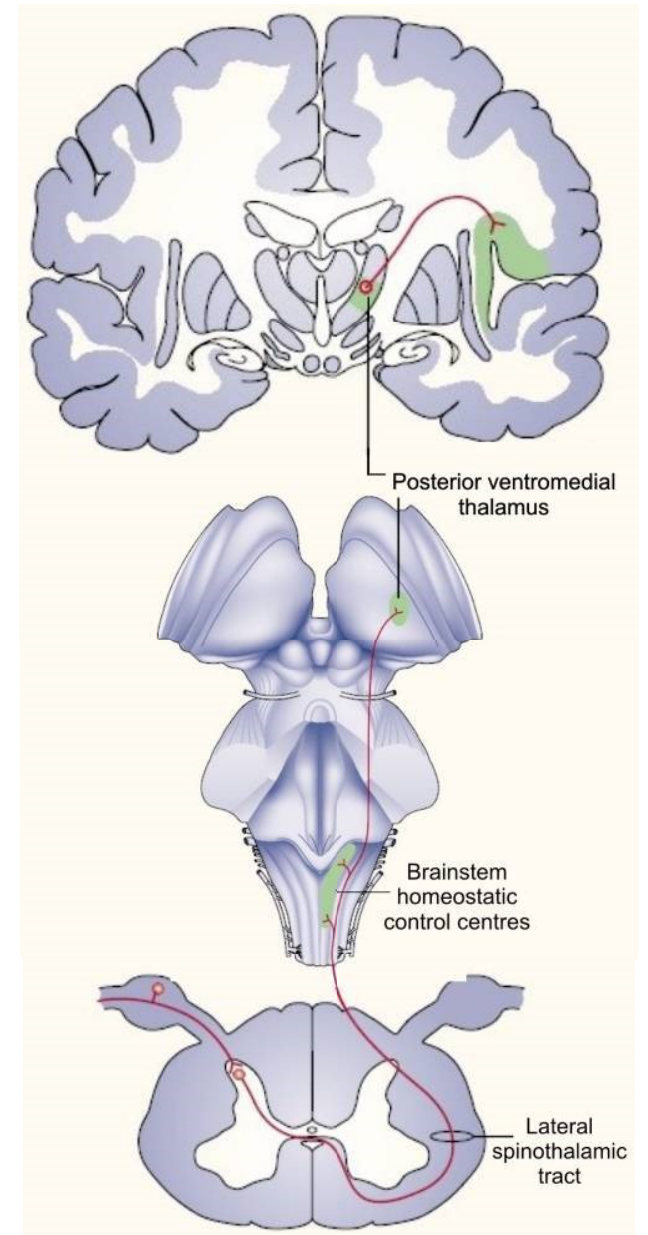


# Interoception

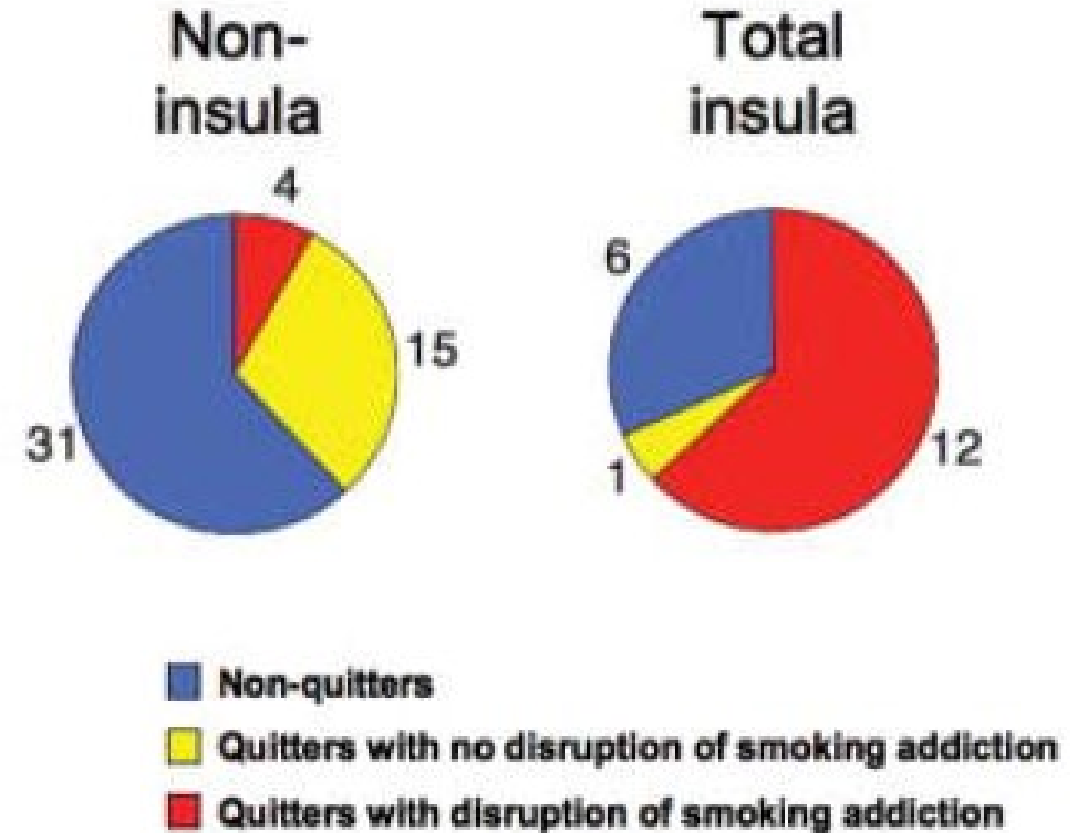
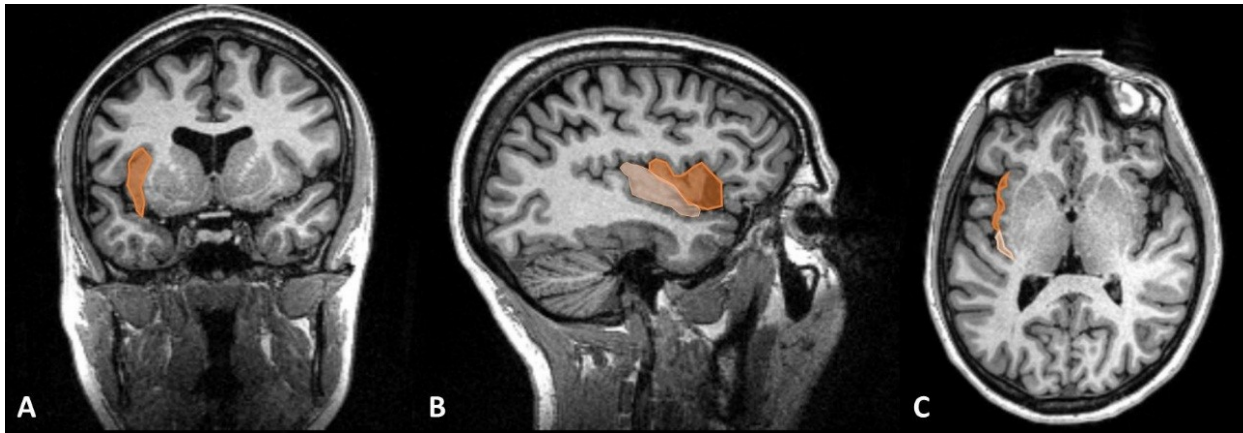
General sensitivity to bodily sensations  
*e.g. heart beat, visceral pain, thirst*

Crucial facet of emotional regulation

Insula, the “interoceptive hub”

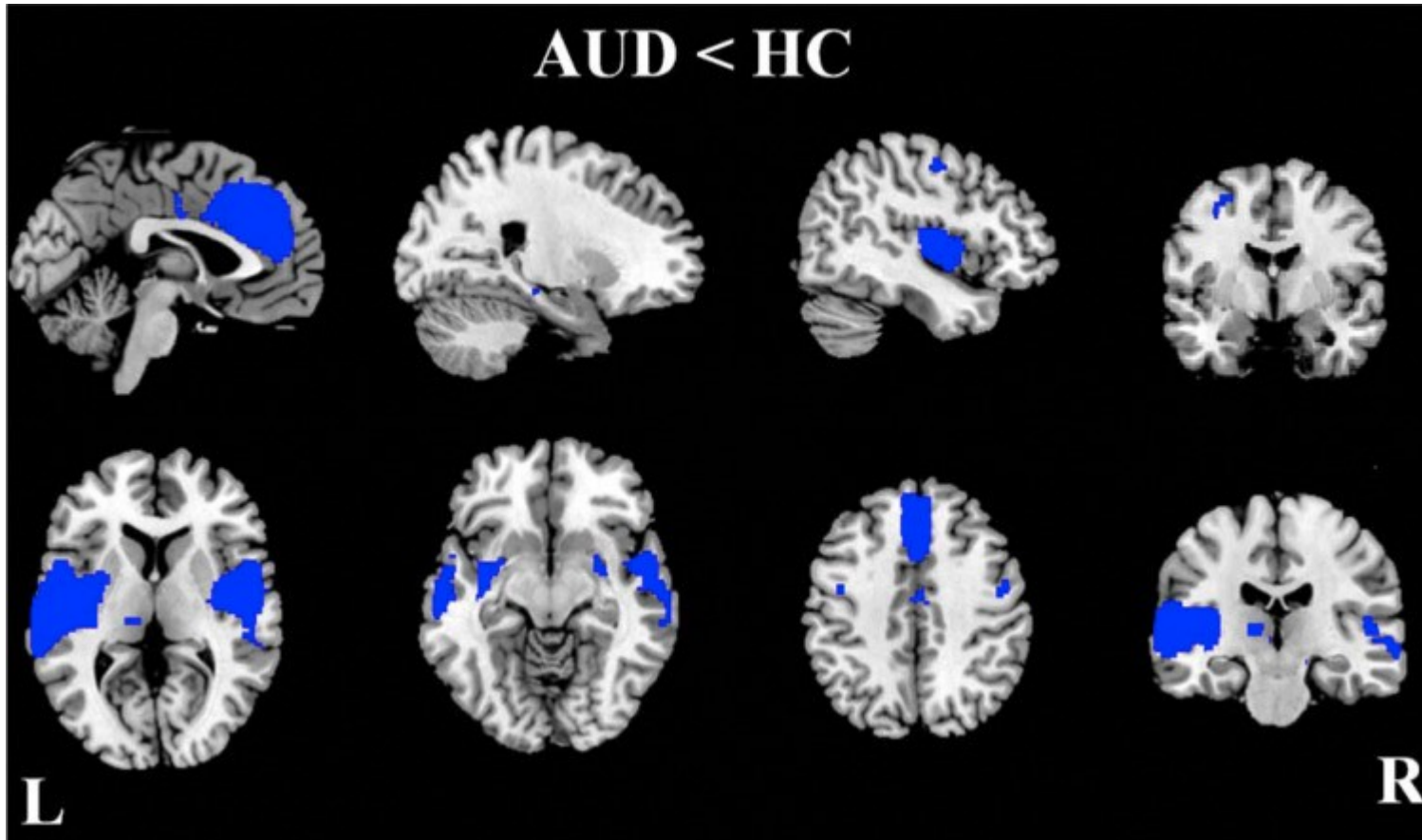


# Addiction and Insula



Insula lesion → Increased likelihood of smoking addiction disruption

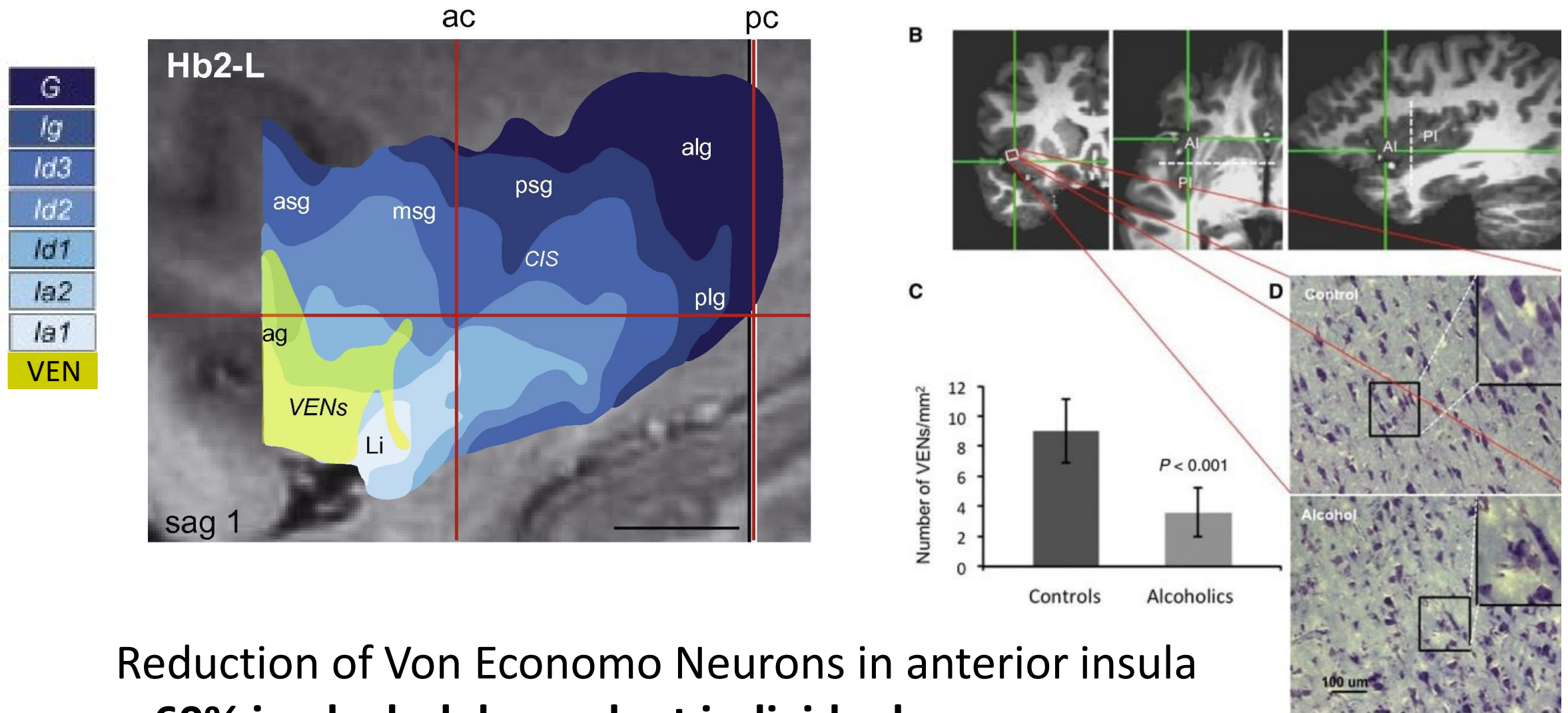
# Insular Volume and Alcohol



a voxel-based meta-analysis

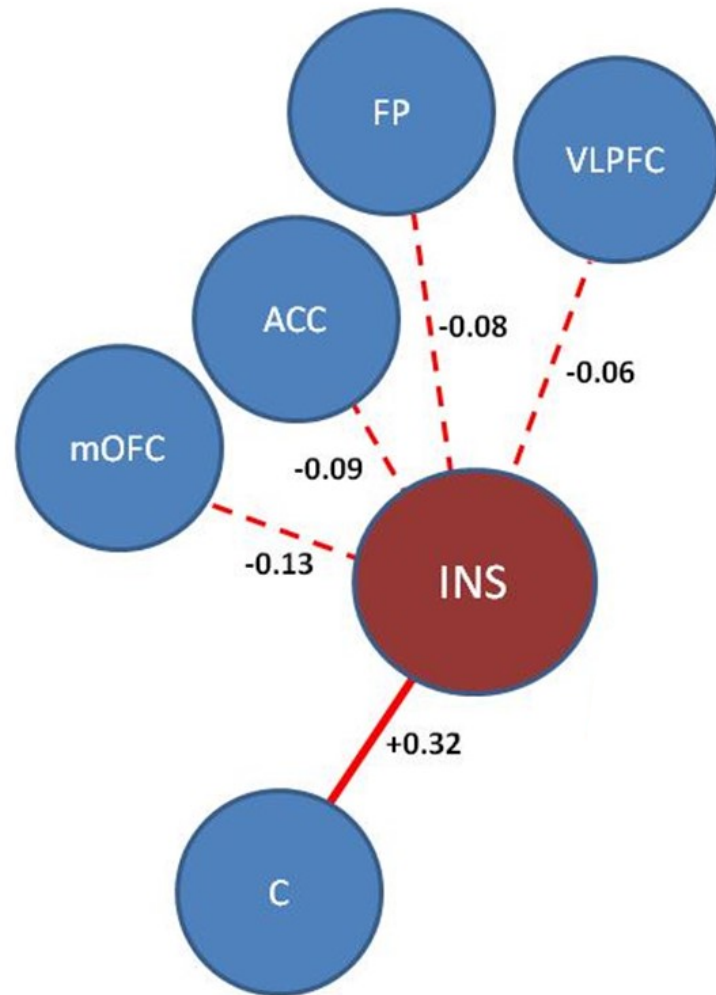
**Cortical and  
subcortical grey  
matter shrinkage of  
the brain**

# Insular Von Economo Neurons and Alcohol



Reduction of Von Economo Neurons in anterior insula  
- **60% in alcohol dependent individuals**

# Insular Connectivity and Alcohol



**Decreased connectivity** between insula and frontal regions in alcoholism

**Reduced white matter integrity** in heavy drinkers correlated with increased insular activation  
=> loss of control over alcohol

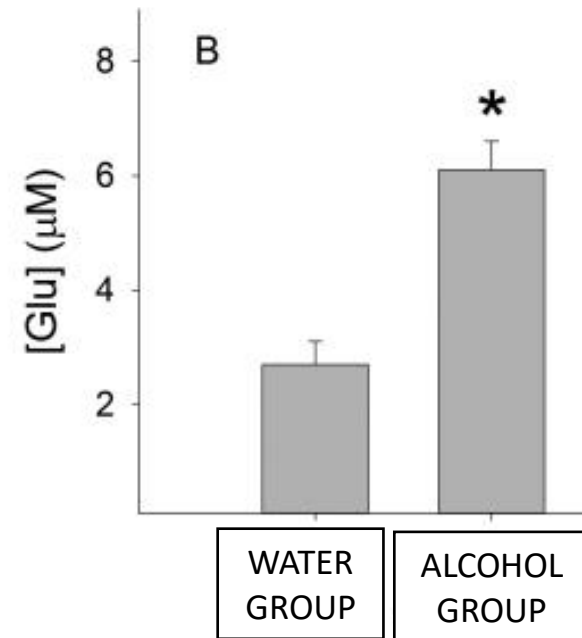
# Insula, Glutamate and Alcohol

Projections on (insular) pyramidal neurons are **glutamatergic** (main excitatory neurotransmitter)

Alcohol **interacts** with glutamatergic neurotransmission

Leading to **excitotoxicity** and neuronal death

Most of studies focused on **frontal or reward-related areas** => what about the **insula**?





# Objective & Hypotheses

## **Measuring insular glutamate concentration, volume and surface parameters in social drinkers**

Insular glutamate concentration will be negatively correlated with social drinking

Insular grey matter volume and cortical folding will be negatively correlated with social drinking

# Methods

## Participants

32 Healthy male social drinkers (age  $M=25.09$  years)  
Weekly units drunk were measured ( $M=24.82$ ,  $SD=18.09$ )

## Questionnaires

-Alcohol Use Disorders Identification Test (AUDIT)  
-Obsessive Compulsive Drinking Scale (OCDS)  
*(e.g. drinking-related thoughts, urges to drink, and the ability to resist those thoughts and urges).*

Asked to abstain from drinking alcohol 24h before the experiment.  
Breathalysed prior the scan



# Methods

## MRI Data acquisition

1.5 Tesla MRI scanner (Siemens Magnetom Avanto)

*T1-weighted images - 3D MP-RAGE sequence*

Repetition Time (TR) = 2730ms

Echo Time (TE) = 3.57ms

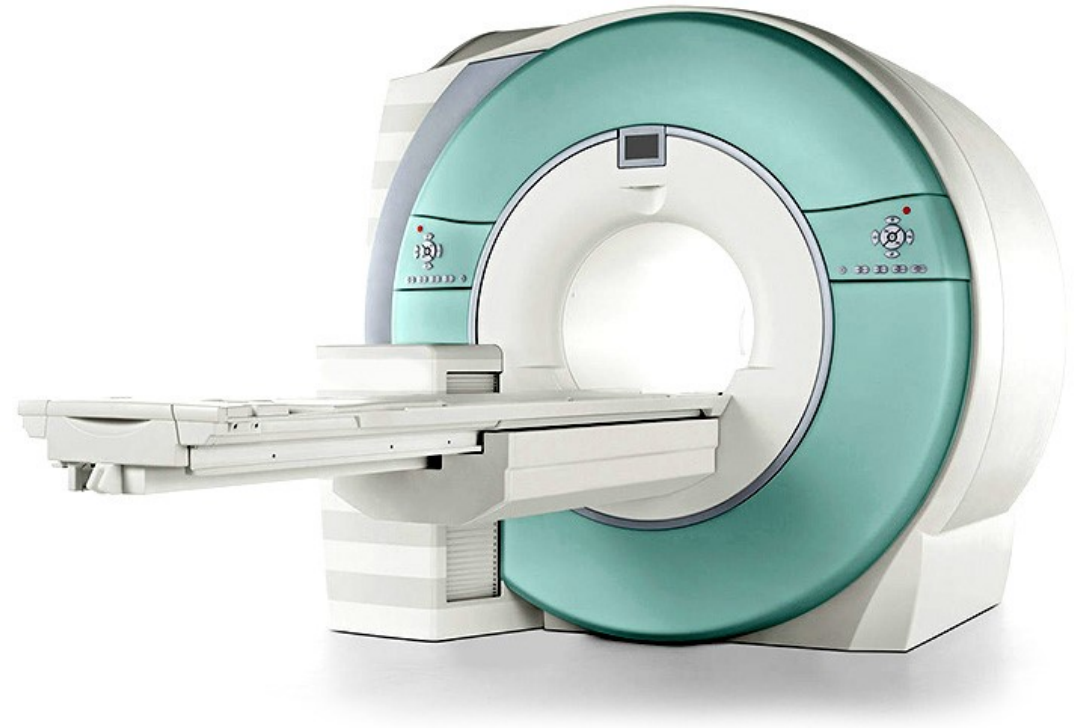
7° Flip angle

Matrix = 256x256

Field of view (FOV) = 256x256mm

1.0 mm isotropic voxel size

192 slices



# Methods

## MRS Data acquisition

### *Single-voxel spectroscopy -Point RESolved Spectroscopy (PRESS) Sequence*

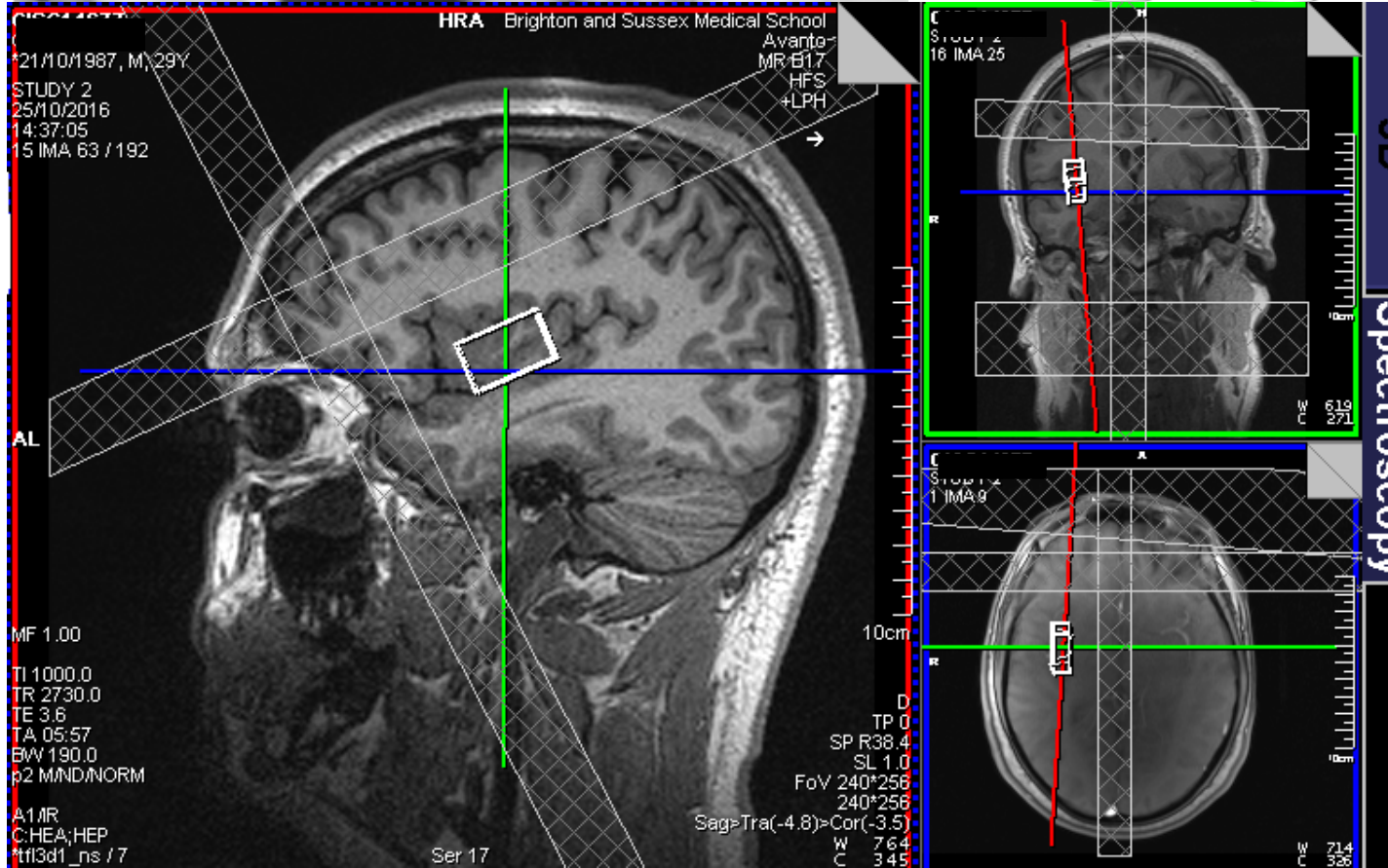
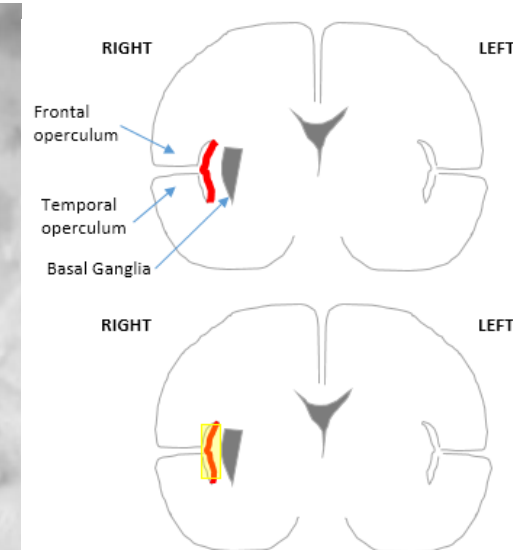
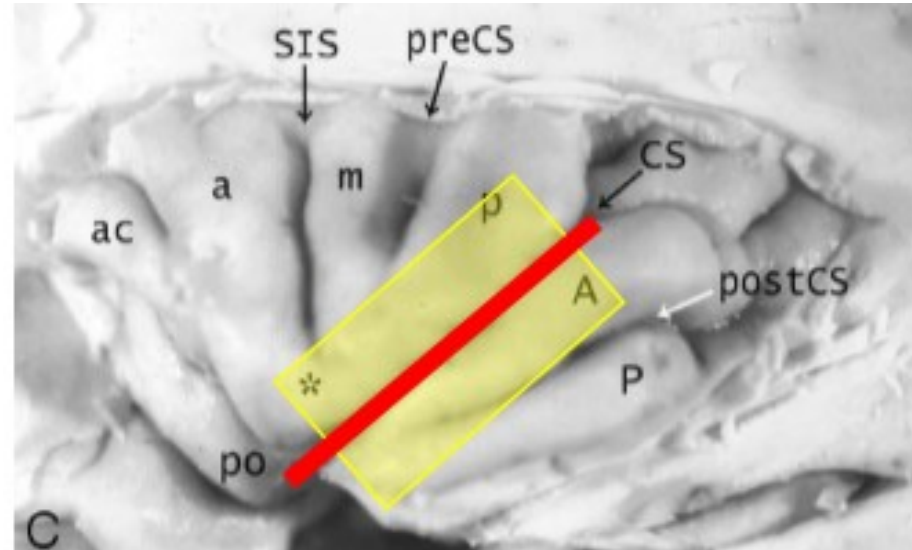
TR=2000ms; TE=40ms; 90° flip angle;  
128 excitations; FOV =16 cm

Voxel size 10x15x25 mm

Right mid-insula

Glutamate + Glutamine (Glx) ;  
Glutamate (Glu) and Glutamine (Gln)

Bottomley et al, 1987; Jang et al., 2005



# Methods

## MRS Data analysis

Creation of a binary mask of the VOI

Resliced to the T1 (SPM12)

T1 was segmented

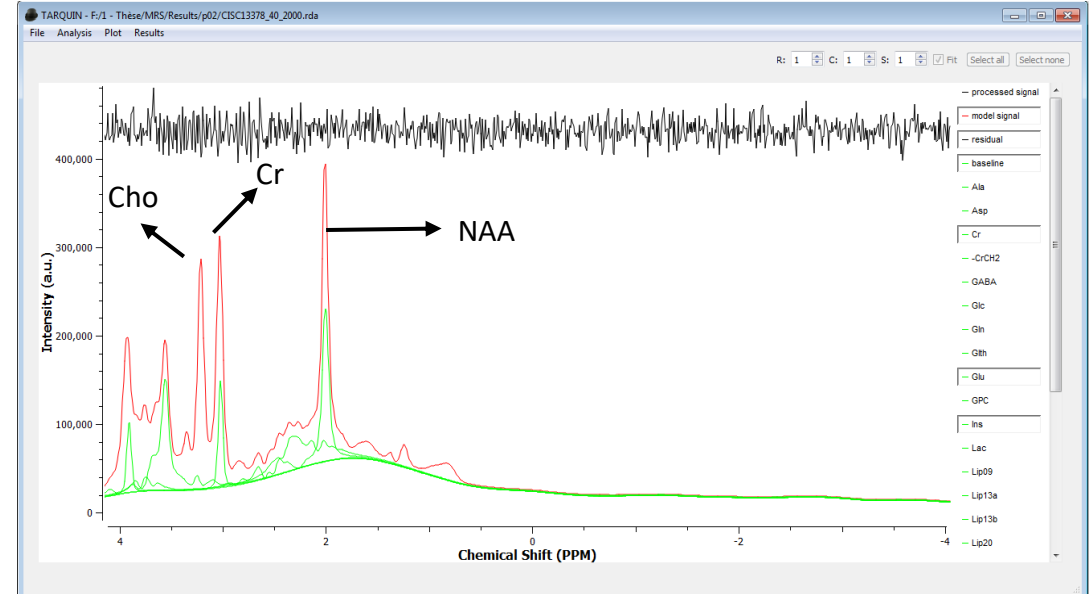
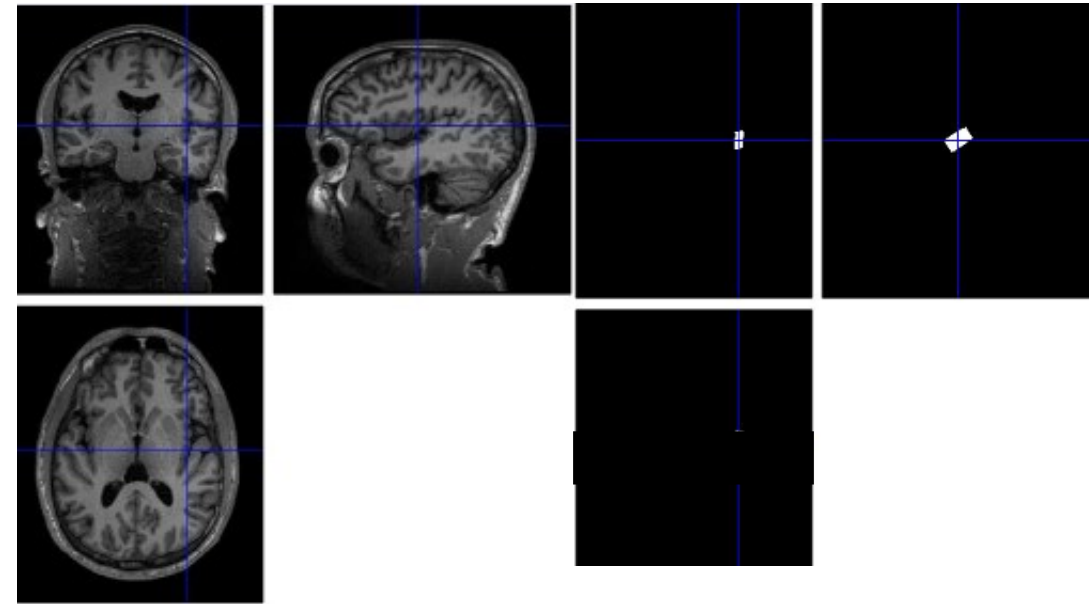
Internal water as reference

Corrections: partial volumes, water attenuation, water concentration, water and metabolites relaxation effects

Tarquin 4.3.7

(water attenuation=1;

water concentration 55510 mM)



# Methods

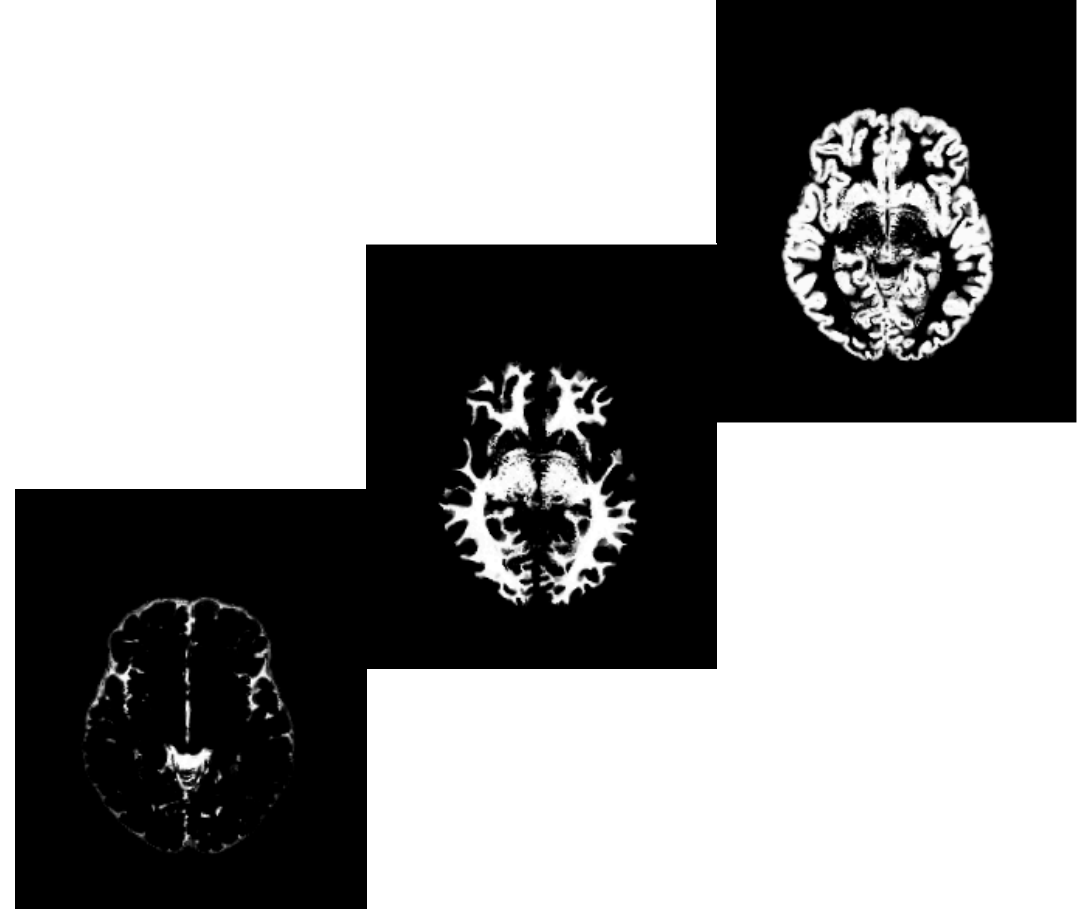
## MRI Data analysis

Voxel and Surface based Morphometry (CAT12)

Region of Interest/ ROI: Extraction

**Grey matter volume** and **cortical gyrification**

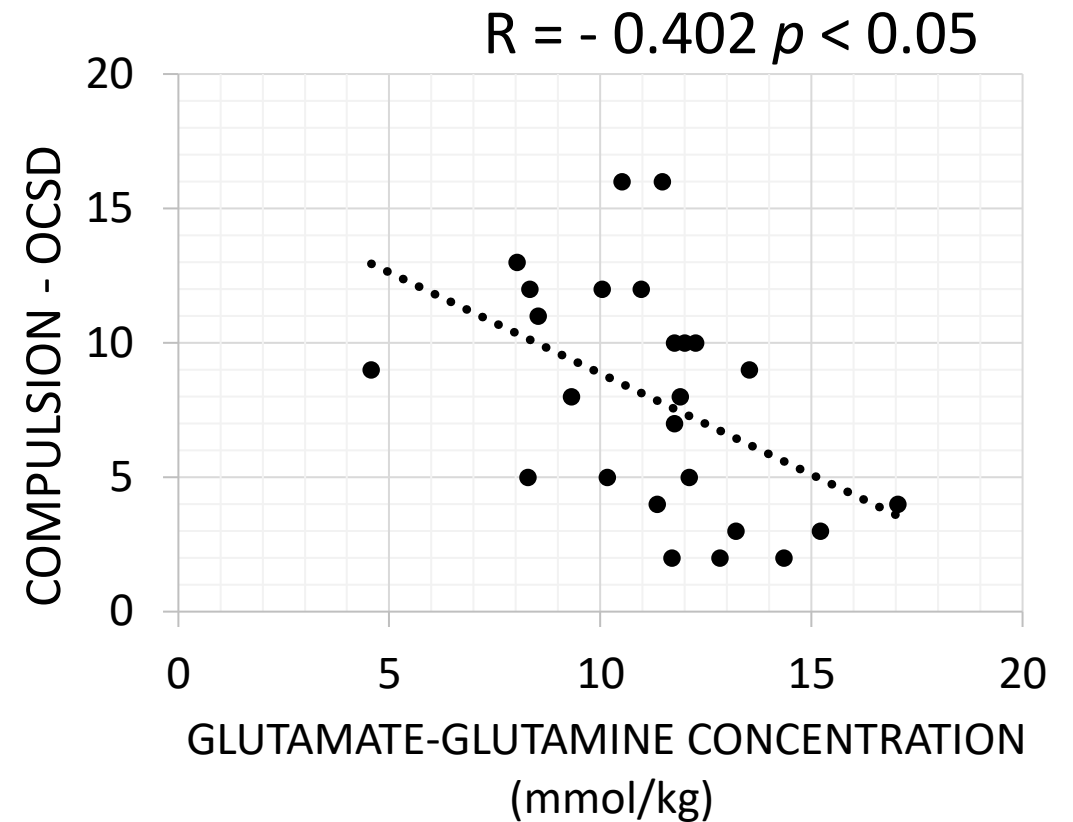
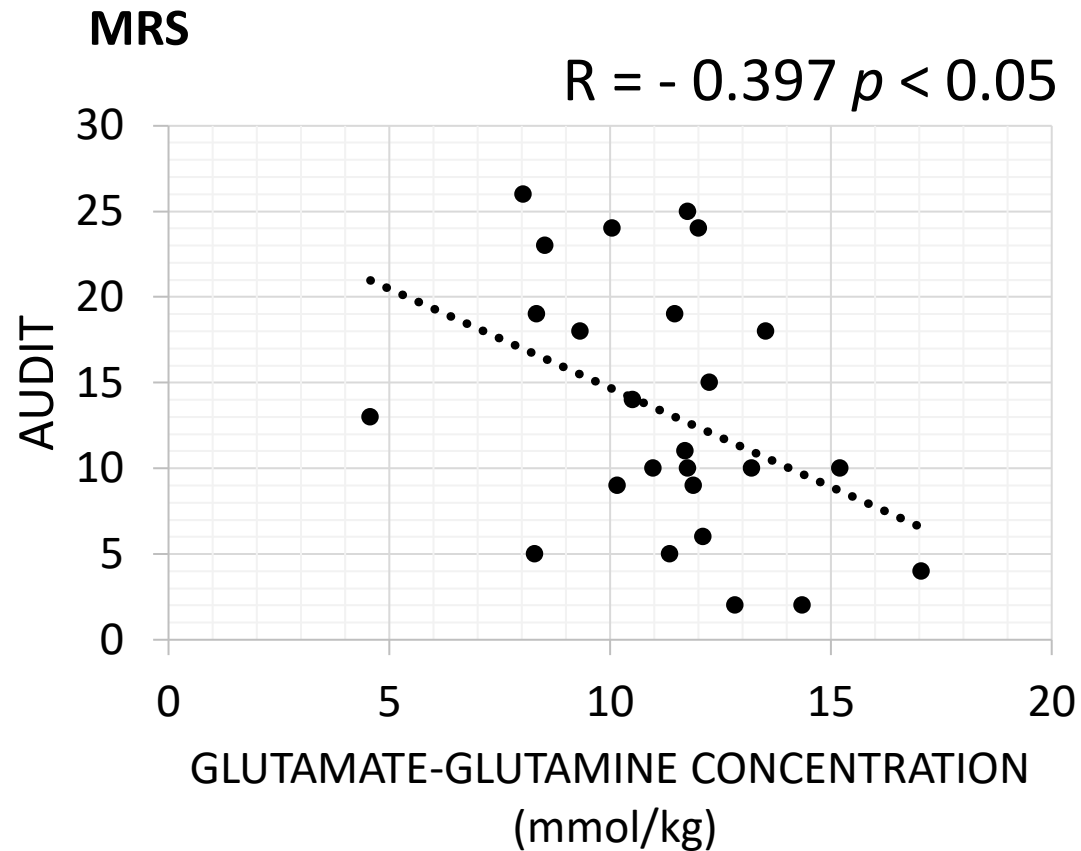
before normalization from right insula



## Statistical Analysis

Correlations and regressions (controlling for age and Intracranial volume) using SPSS 24

# Results



**Alcohol use disorder severity and alcohol compulsions are related to reduction of insular glutamate and glutamine concentrations**

# Results

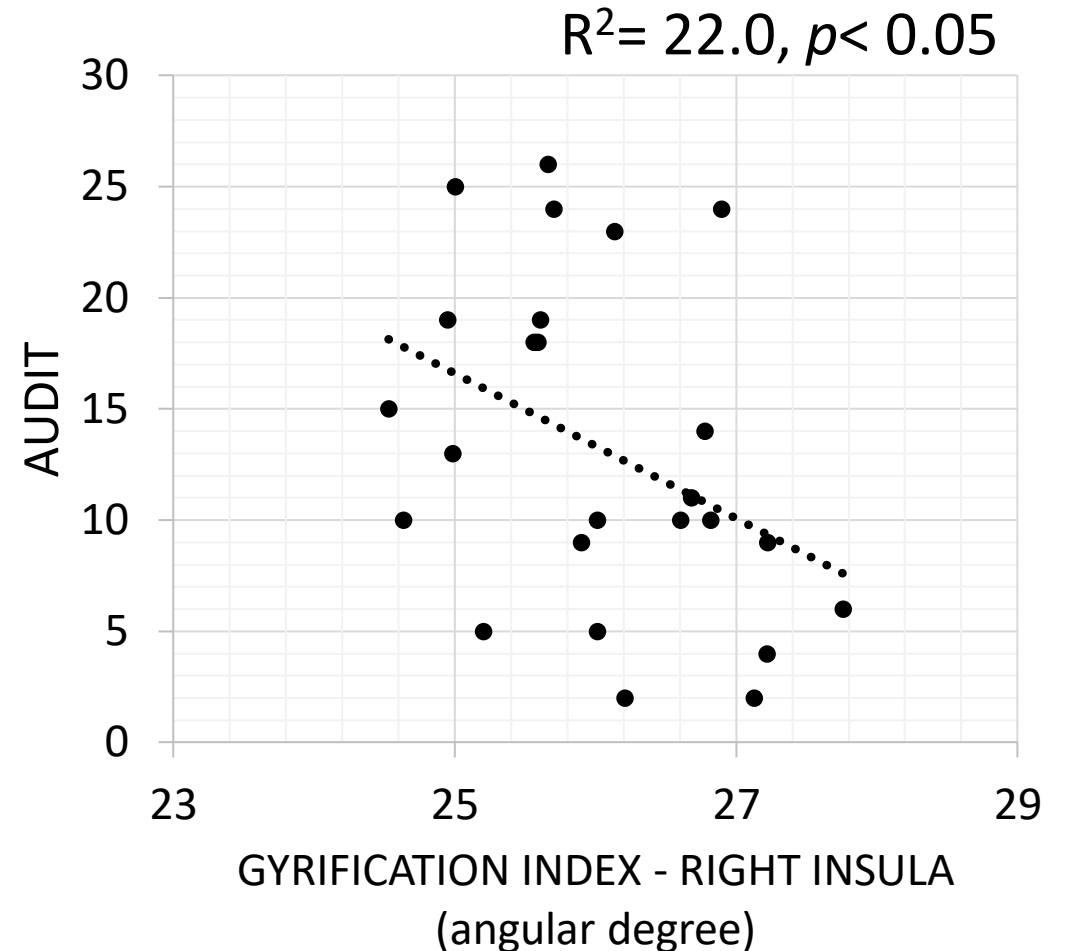
**SBM**

ROI – right insula

Alcohol use disorder severity was predicting reduced gyrification index in the right insula.

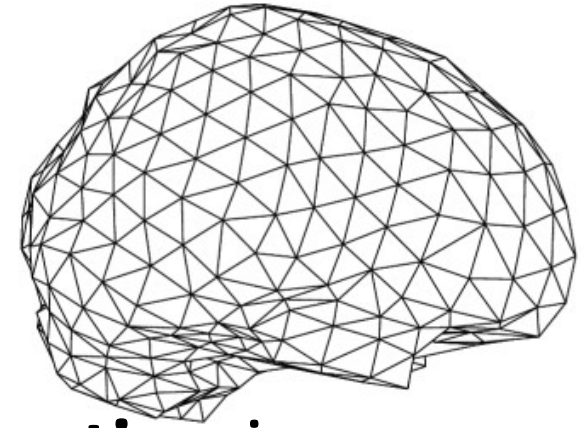
Severity of alcohol use disorder is associated with a **decreased cortical folding** in the right insula.

**No VBM results**





# Conclusion



**Reduced insular glutamate and glutamine concentration in heavy drinkers (mainly driven by glutamate).**

**Insular gyrification reduction but no insular volume reduction**

Atrophy => metabolite reduction => interoception impairment

Subtle changes?

Is cortical gyrification a more sensitive measure of early alcohol-associated decline in grey matter?

# Impact

First study looking at the interoceptive component of alcohol seeking

Integrity of insular cortex

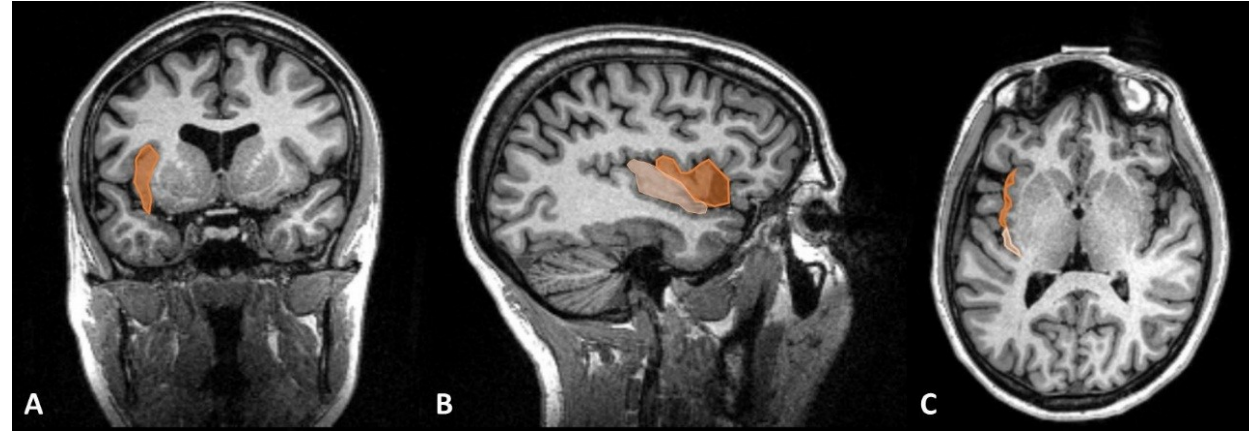
In social drinkers

Using multimodal neuroimaging



# Implication

The insula :



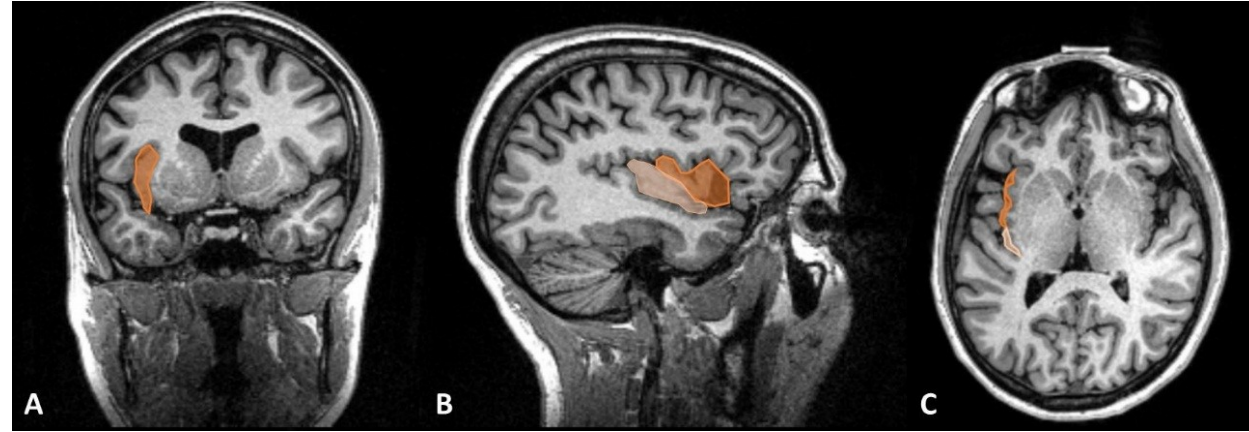
- Central role in the interoceptive component of drug seeking
- Impaired in alcohol and drug use disorders

**IF**

**insular disruption ↔ compulsion ↔ relapse**

# Implication

The insula :



- Central role in the interoceptive component of drug seeking
- Impaired in alcohol and drug use disorders

**IF**

**insular disruption ↔ compulsion ↔ relapse**

# Thank you for your attention!

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Thanks also to everyone  
involved

