



Spinal FMRI and Physiological Noise Correction

Jonathan Brooks

CRiCBristol, University of Bristol



What's covered in this talk

- What is physiological noise?
- Why does it matter?
- How do I measure it?
- What to do with the data....
- Is it worth the hassle?
- Spinal fMRI

A sagittal MRI scan of a human spine, showing the vertebrae and intervertebral discs in grayscale. The spine is oriented vertically, with the head at the top and the pelvis at the bottom.

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What is physiological noise?

- Sources: cardiac, respiratory, movement
- Importance: at higher field strength, dominant source of noise ($\sigma_{\text{physio}} \propto \text{field strength}$)
- Effects: additive, will confound signal detection particularly areas of low SNR, or regions with large CSF spaces, near large vessels etc.

Glover et al. MRM (2000)

Kruger & Glover, MRM (2001)

Triantafyllou et al. NeuroImage (2005/2006/2011)

A sagittal MRI scan of the human spine, showing the vertebrae and intervertebral discs. The image is in grayscale and occupies the left side of the frame.

What does it look like

What does it look like



What does it look like





Sources of noise

$$\sigma = \sqrt{\sigma_s^2 + \sigma_T^2 + \sigma_P^2}$$

- Thermal/scanner
- Cardiac
- Respiratory
- Autonomic

Kruger & Glover, MRM (2008)

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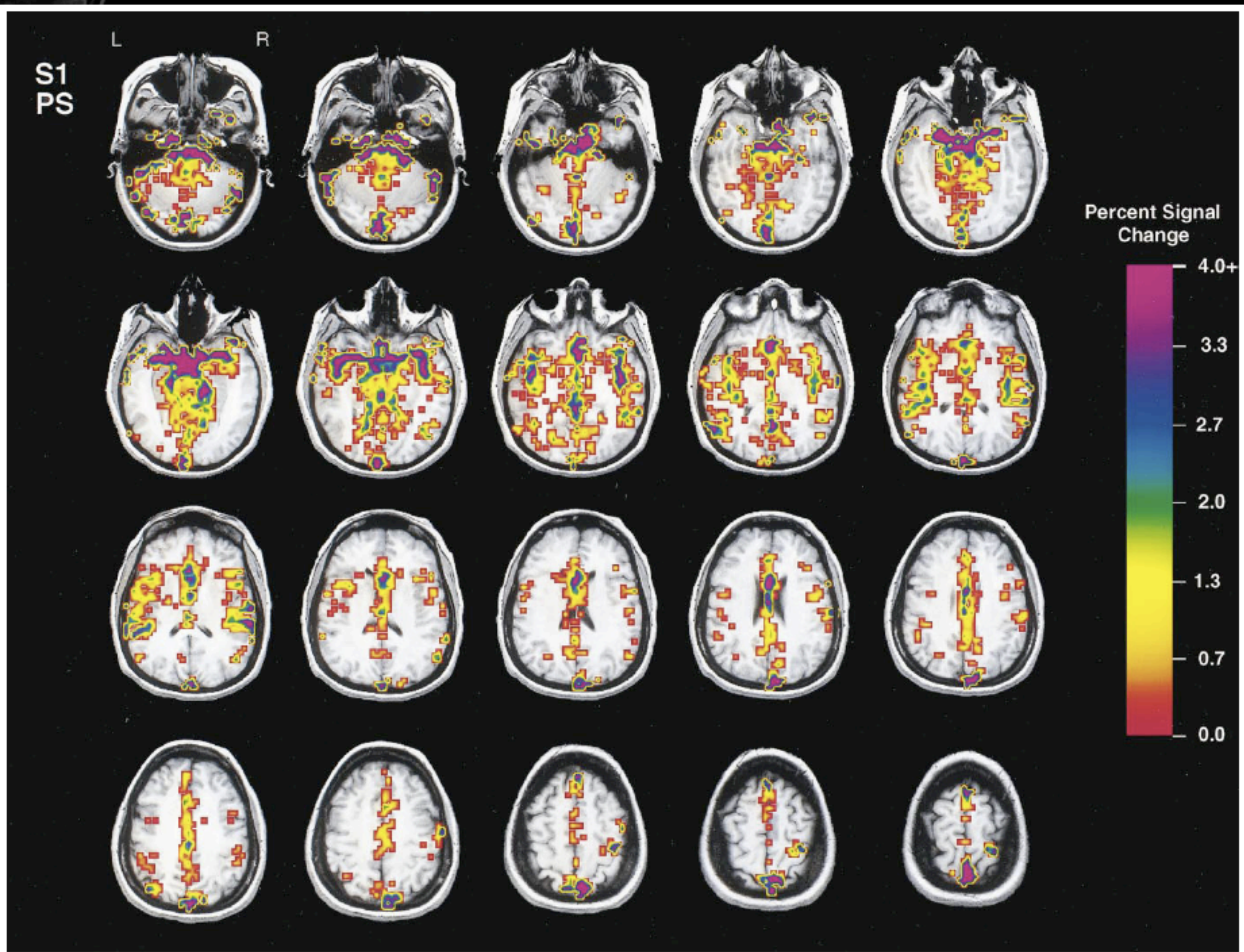
A sagittal MRI scan of the spine, showing the vertebrae and intervertebral discs. The image is in grayscale and occupies the left side of the slide.

CARDIAC

- Pulsatile movement (arteries, capillaries, brain tissue)
- BOLD-like effects?
- Increased CBV, but fixed cranial capacity
⇒ movement of CSF



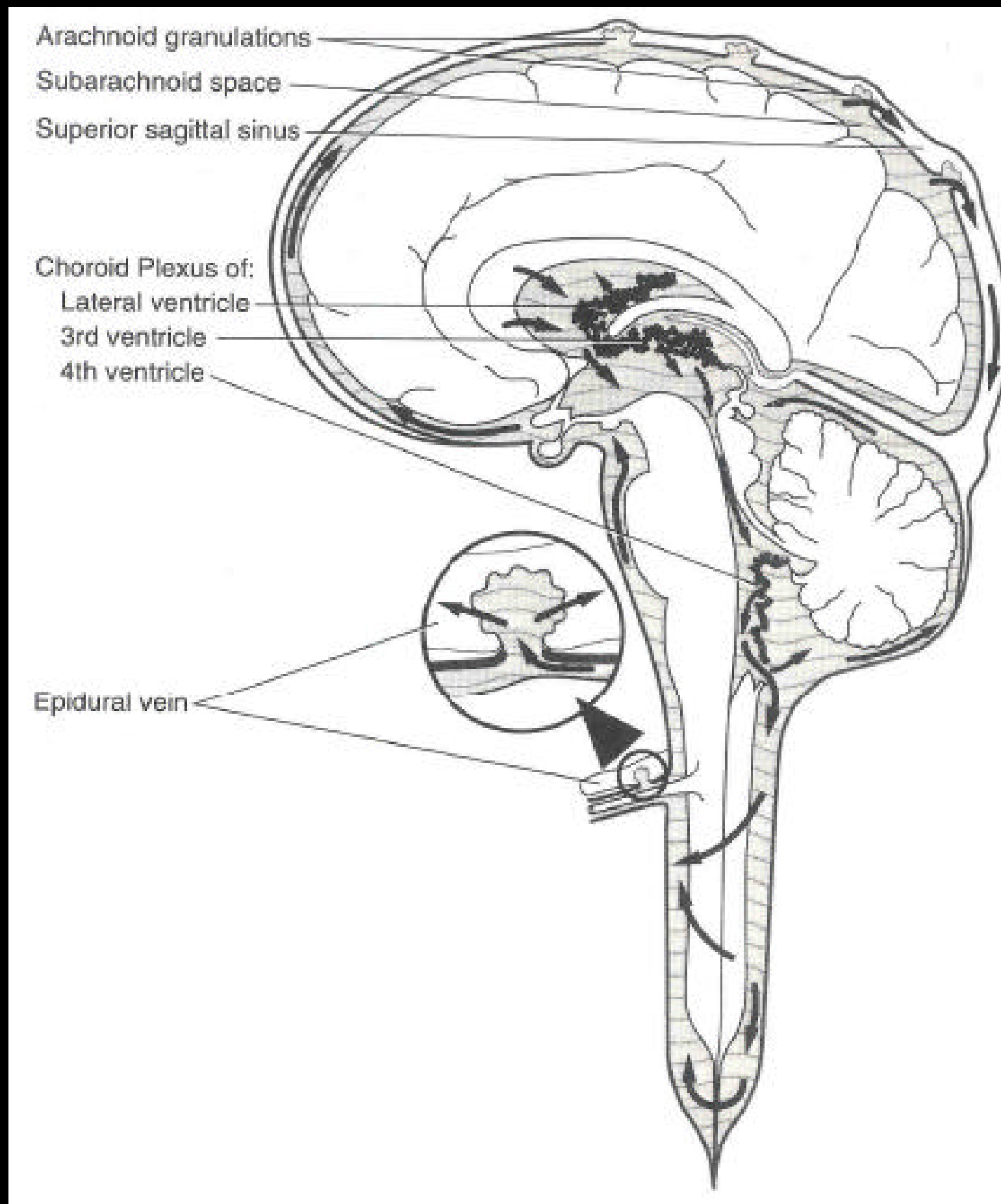
Dagli et al, NeuroImage (1999)



Dagli et al, NeuroImage (1999)

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Kruger & Glover, MRM (2008)

- Thermal/scanner
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- **Respiratory**
- Autonomic



RESPIRATORY

- B_0 susceptibility - changing volume of air and position of chest
- Resting fluctuations in rate and depth of breathing can produce systemic change in PaCO_2 , producing vasodilatory (BOLD-like effects)

RESPIRATORY

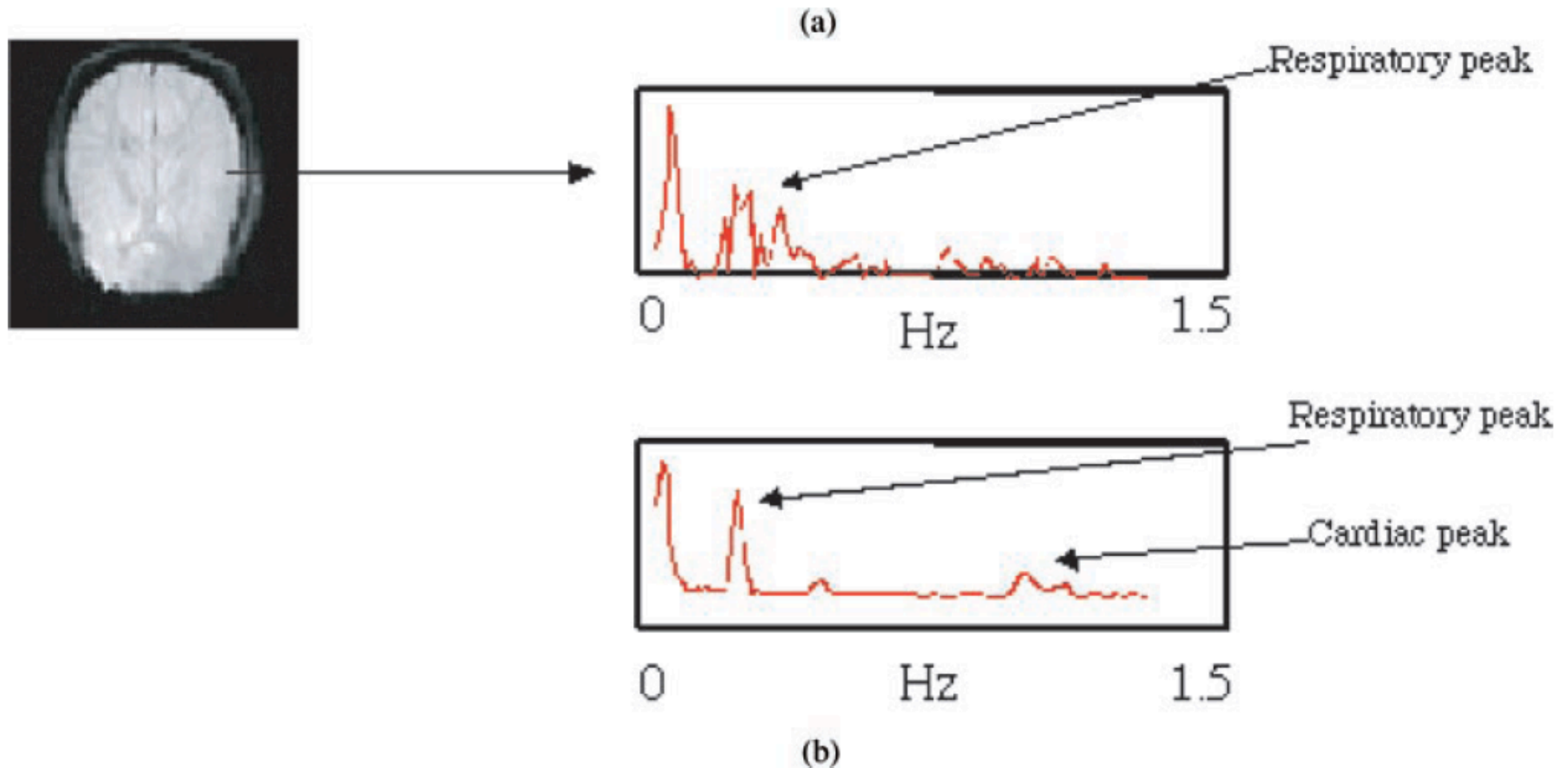


Figure 1. (a) The power spectrum of the signal from a single pixel in grey matter. (b) Averaged power spectrum for pixels across the whole brain.

RESPIRATORY

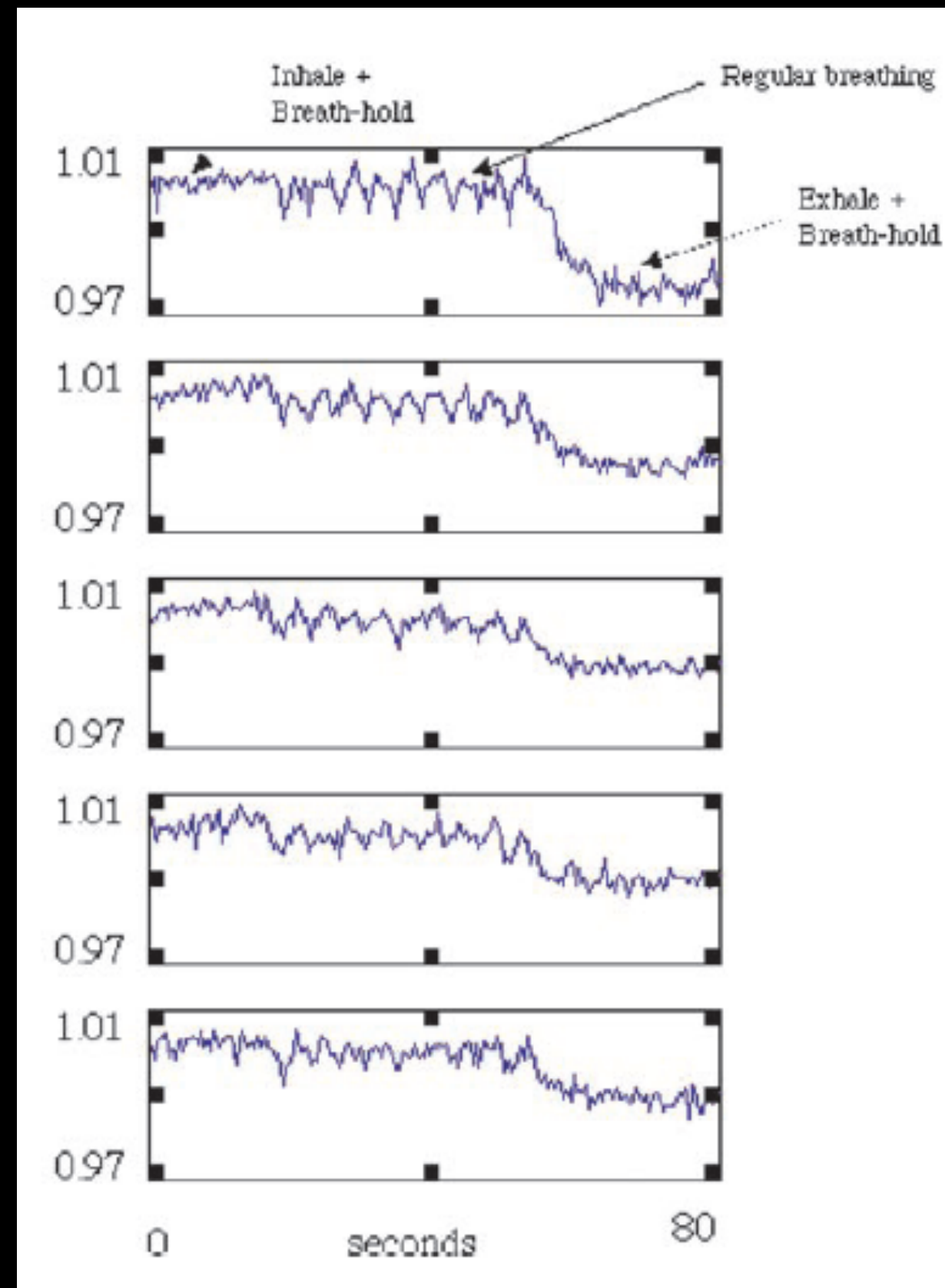
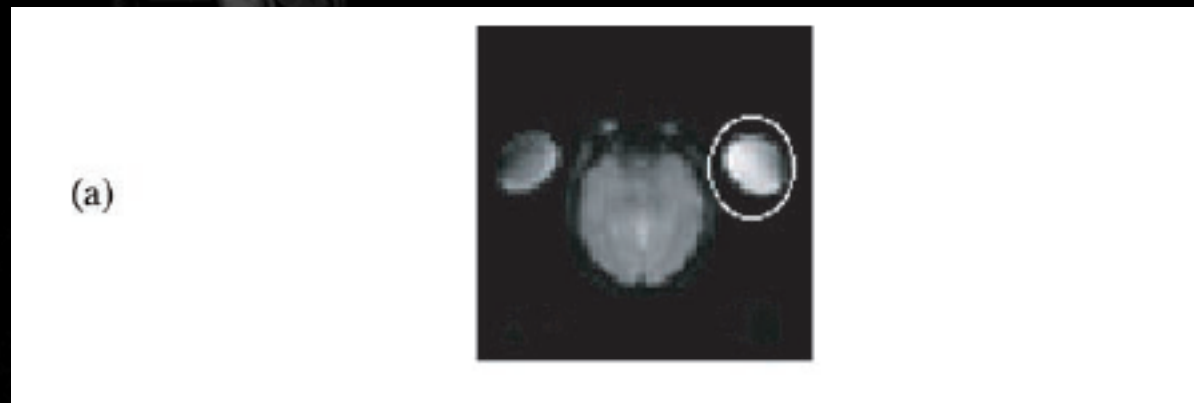
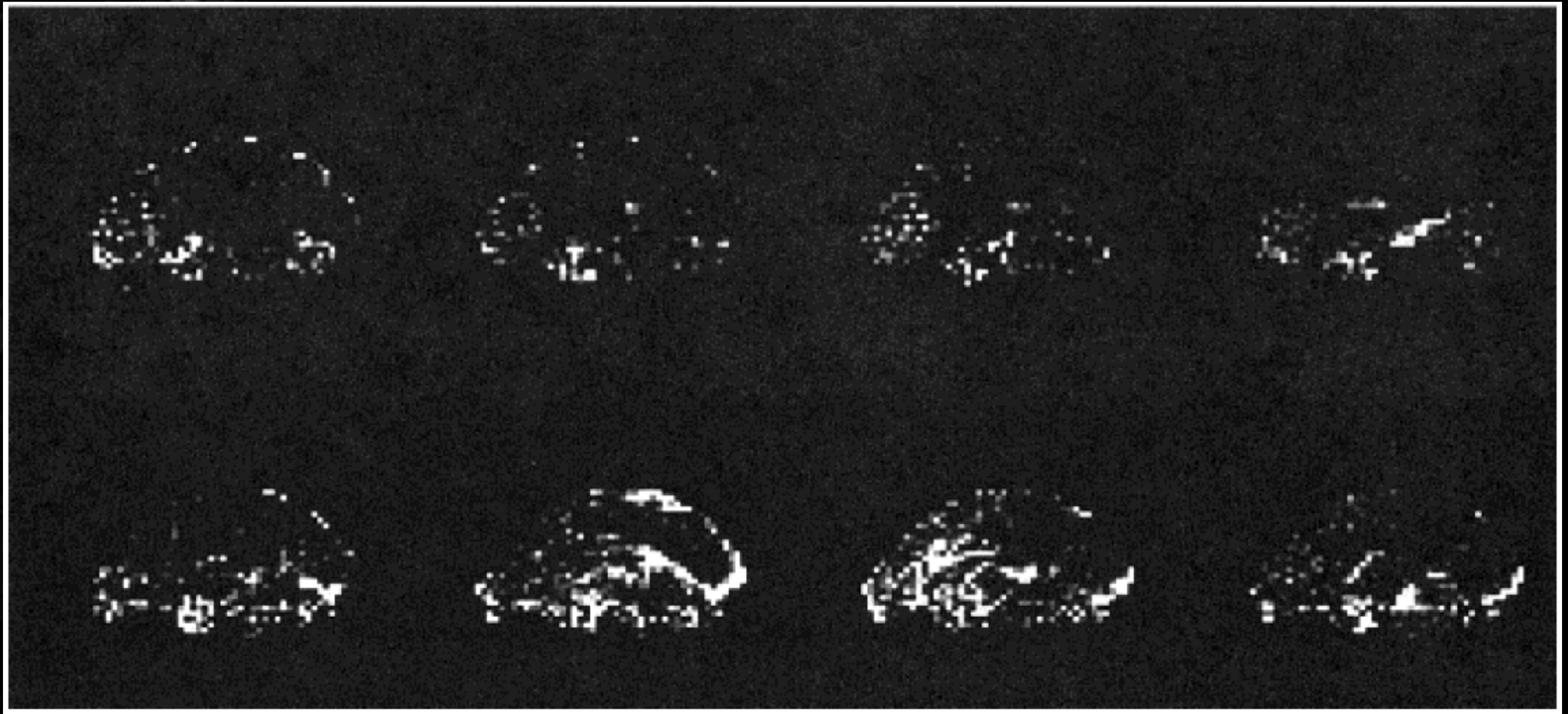


Figure 2. (a) Axial echo-planar brain image acquired with two bottles containing 0.01 mM copper sulfate attached to the head coil. (b) MR signal from a region of interest (10 pixels) in the bottle shown in figure 2(a). During images 1–40, the subject held his breath after exhaling completely, during images 40–140 normal breathing was resumed and during images 140–200 the subject held his breath at full inspiration.

Raj et al, Phys Med Biol (2001)

RESPIRATORY



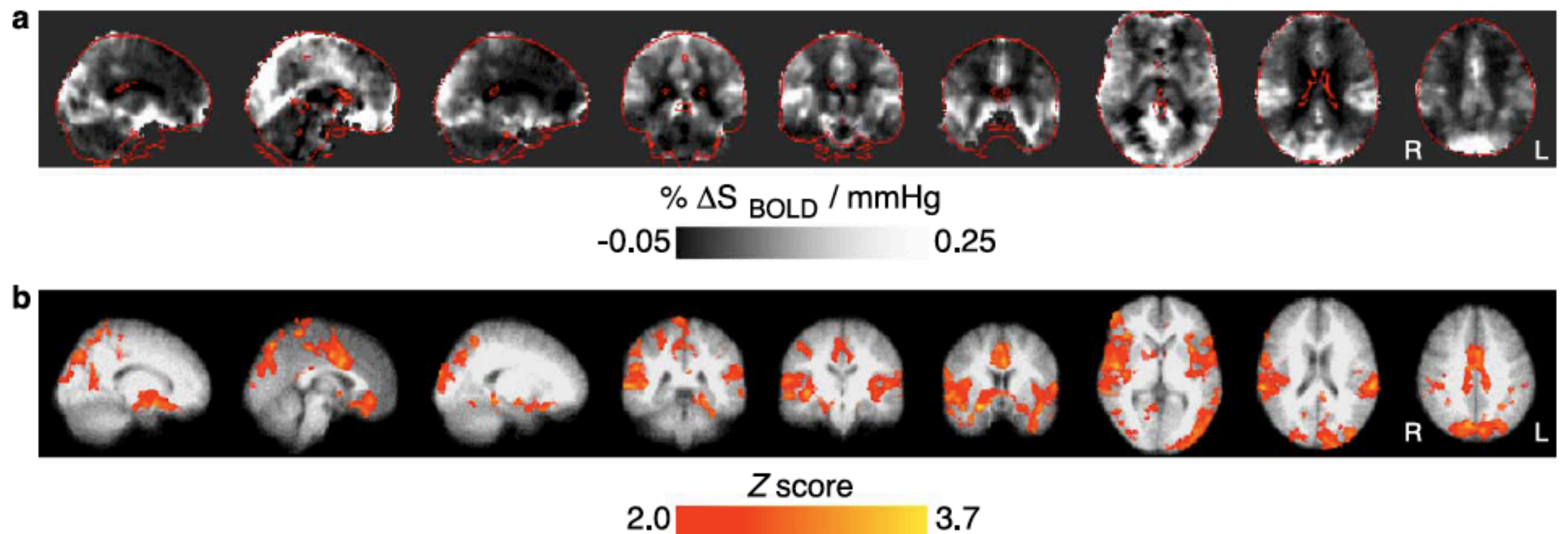
Frank et al, MRM (2001)



RESPIRATORY

- B_0 susceptibility - changing volume of air and position of chest
- Resting fluctuations in rate and depth of breathing can produce systemic change in PaCO_2 , producing vasodilatory (BOLD-like effects)

RESPIRATORY



Wise et al, NeuroImage (2004)



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Why does it matter?

- fMRI relies on principle of pure insertion
- Physiological noise creates time varying signals unrelated to stimulation
- I.e. model does not fit data properly:
 - reduced accuracy of parameter estimates
 - decreases significance
 - increased numbers required to show effect at group level

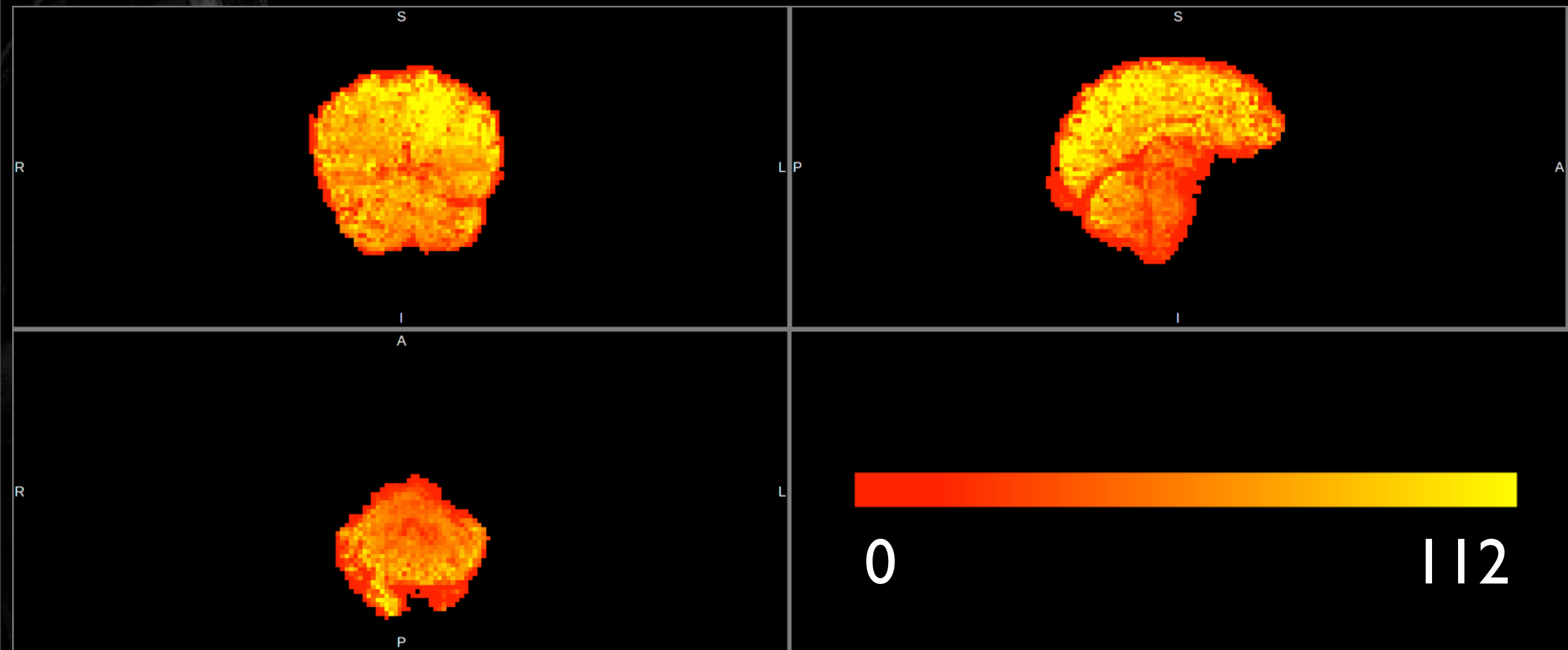
Temporal Signal to Noise Ratio (TSNR)

- Quantity measures intrinsic quality of data
- High signal, and low variability
- Need resting data

$$TSNR = \frac{\text{temporal mean } (T_{mean})}{\text{temporal standard deviation } (T_{std})}$$

Parrish et al (2000) Impact of signal-to-noise on functional MRI.
Magn Reson Med 44:925-932.

TSNR



Murphy et al (2007) How long to scan? The relationship between fMRI temporal signal to noise ratio and necessary scan duration. NeuroImage 34:565-574

A sagittal MRI scan of a human spine, showing the vertebrae and intervertebral discs. The image is in grayscale and occupies the left side of the slide.

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SETUP



SETUP



SETUP



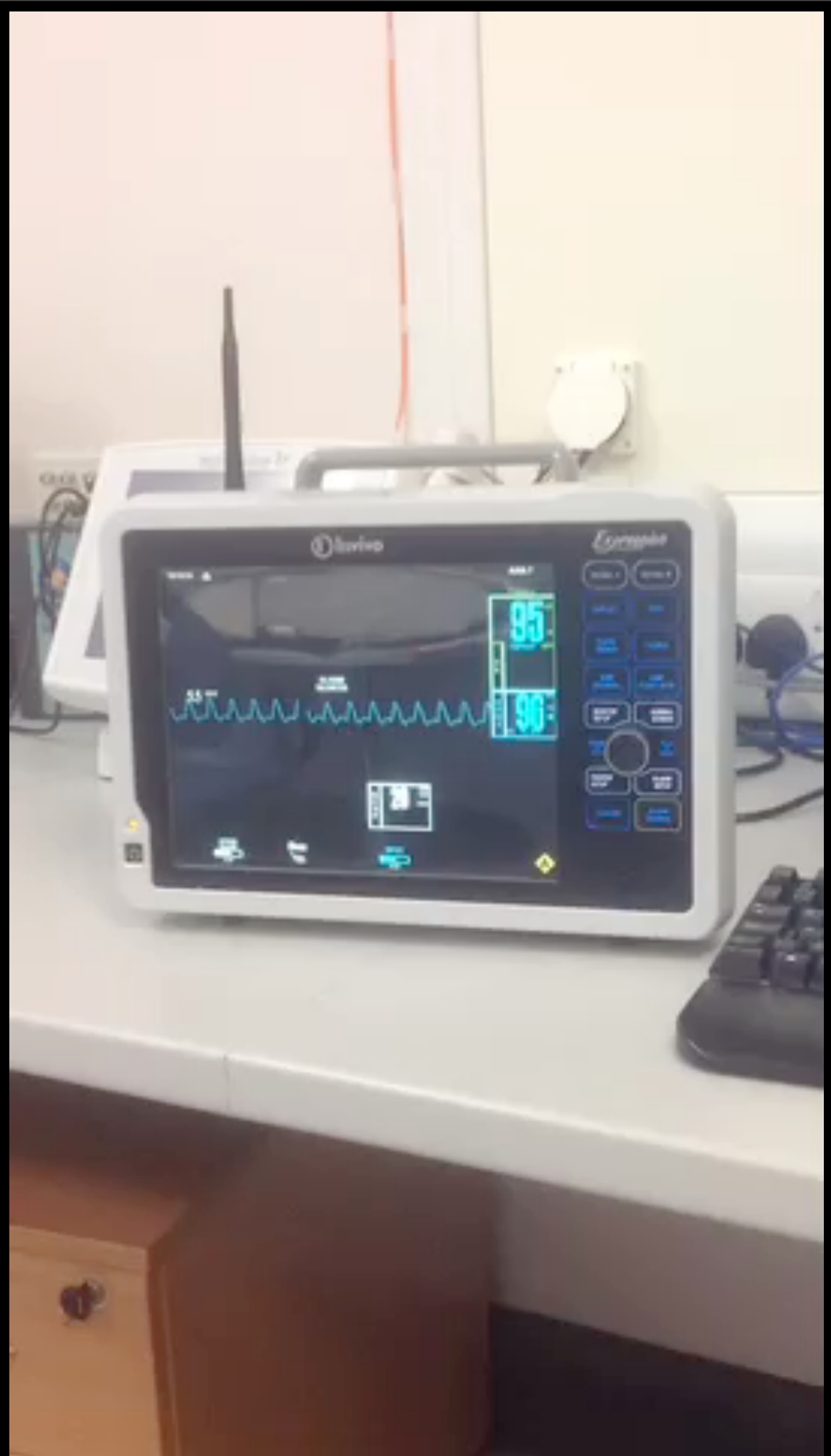
SETUP

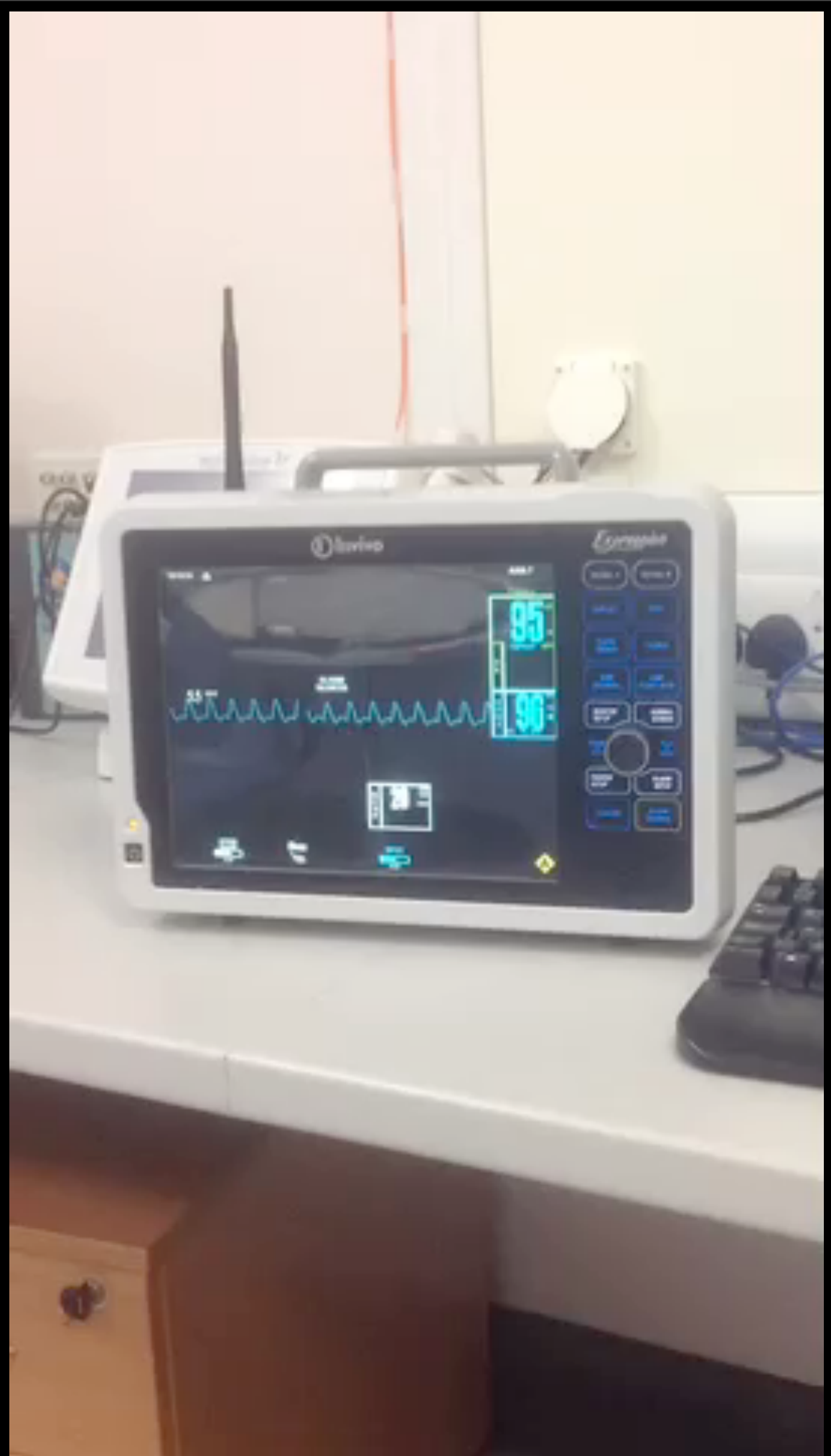


SETUP











What's covered in this talk

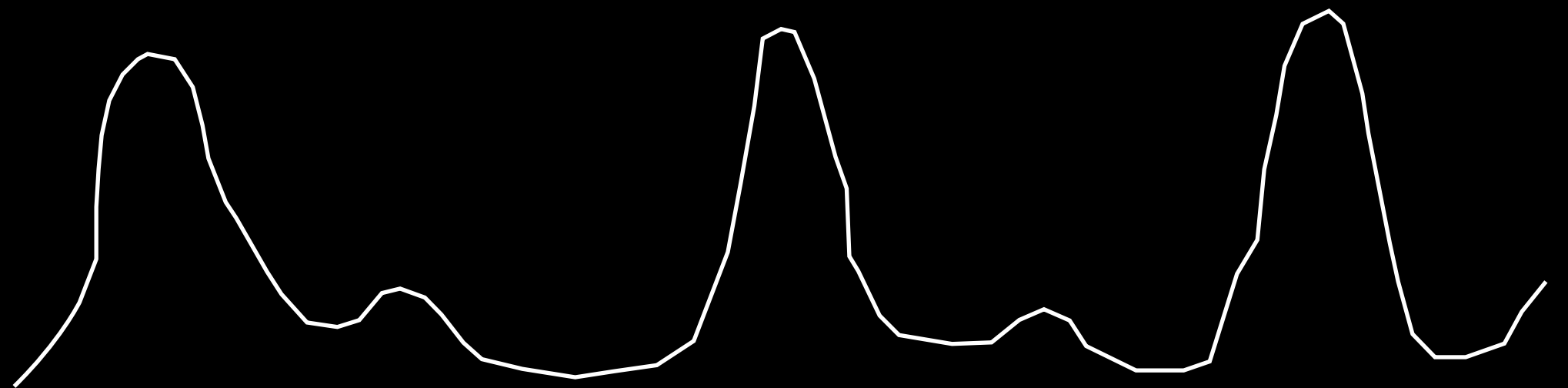
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CARDIAC PHASE



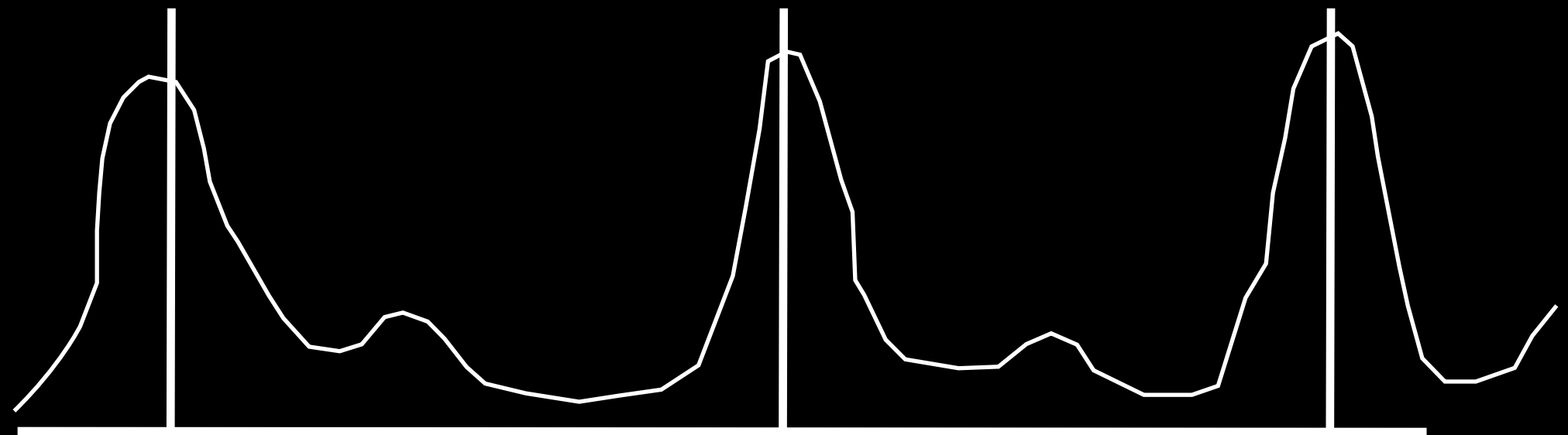
Glover et al, MRM (2000)
Brooks et al, NeuroImage (2008)

CARDIAC PHASE



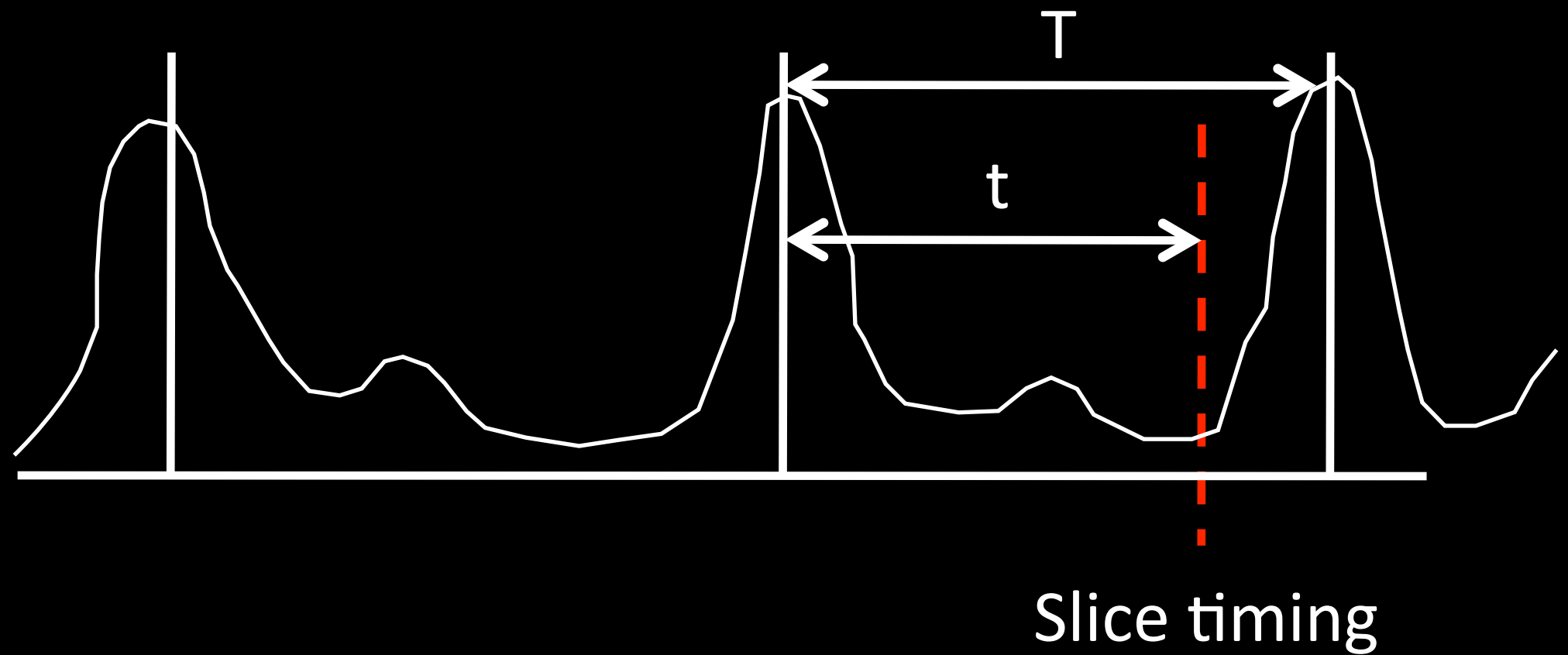
Glover et al, MRM (2000)
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CARDIAC PHASE



Glover et al, MRM (2000)
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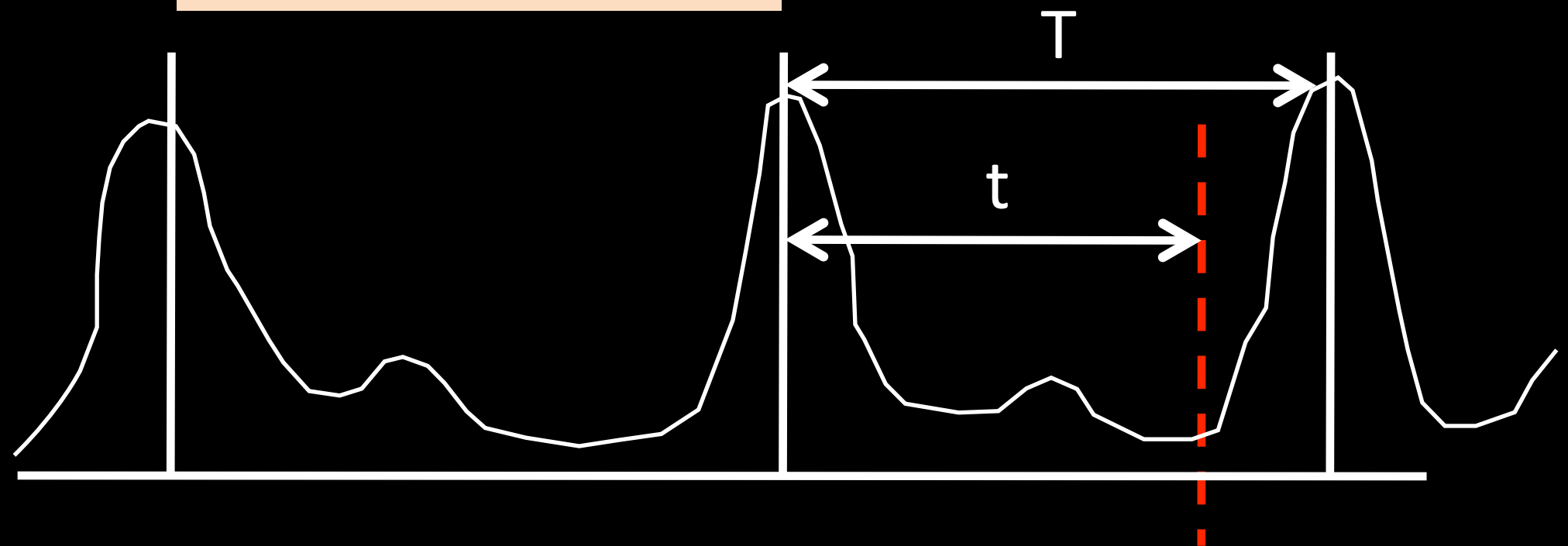
CARDIAC PHASE



Glover et al, MRM (2000)
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CARDIAC PHASE

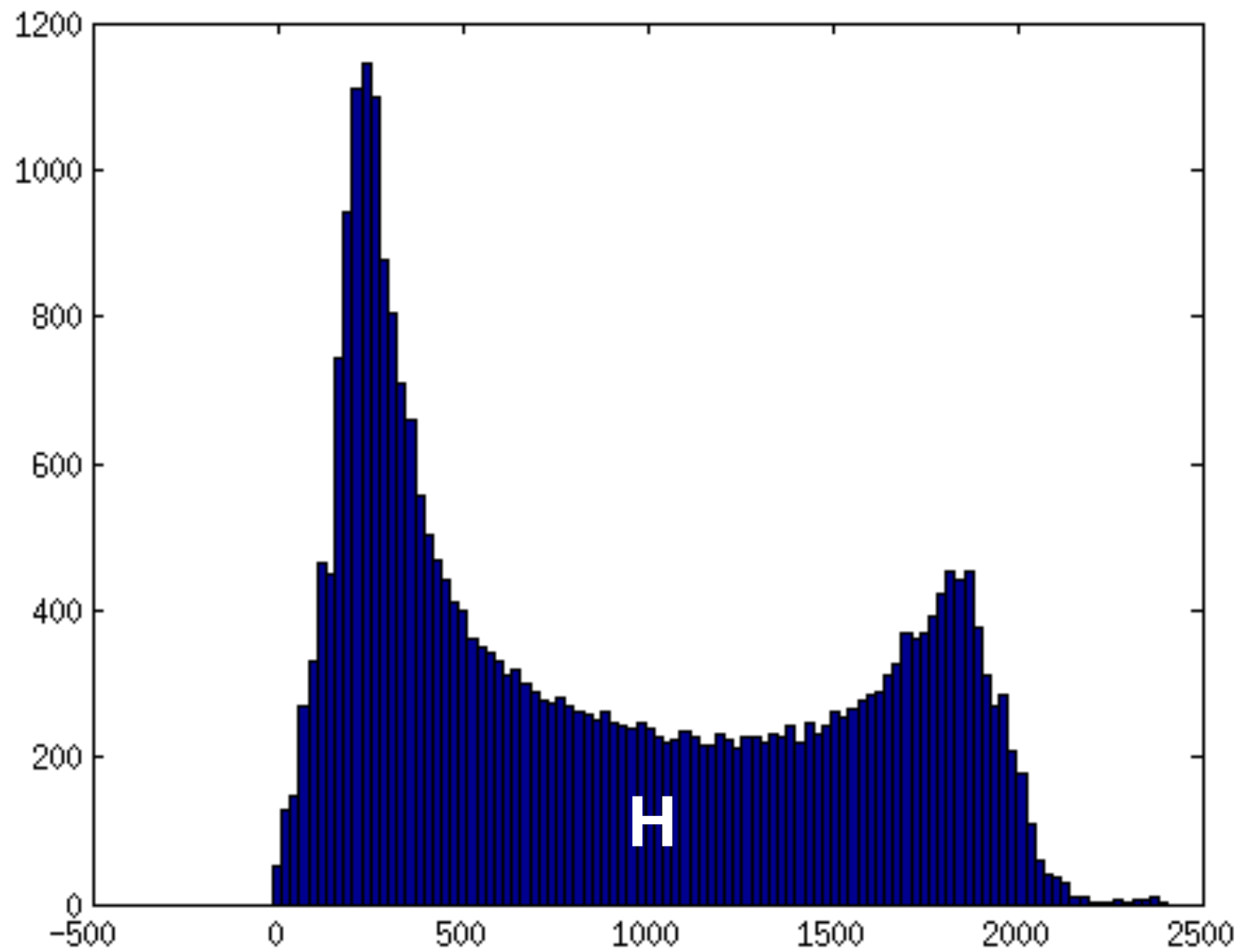
$$\theta_c = 2\pi * (t/T)$$



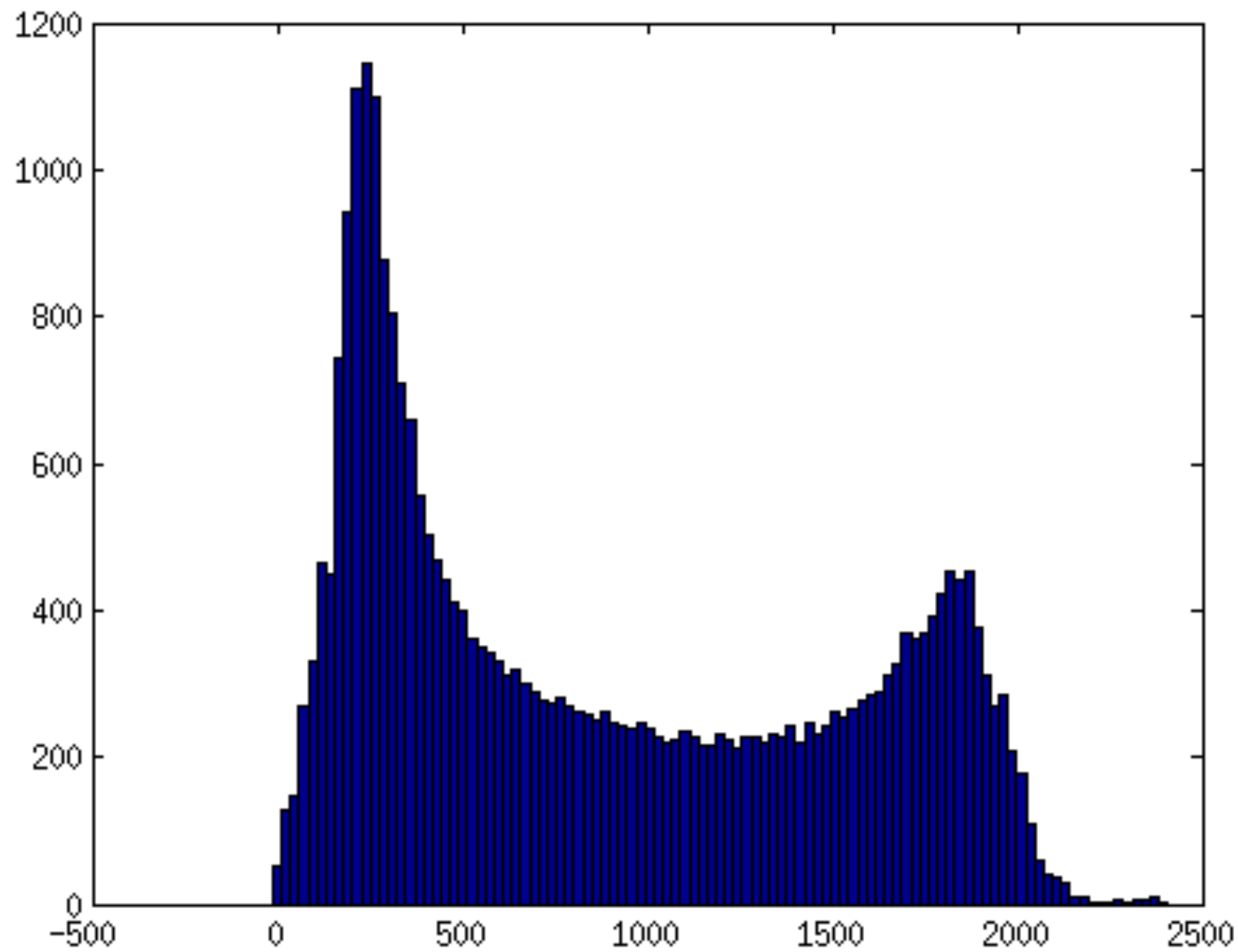
Slice timing

Glover et al, MRM (2000)
Brooks et al, NeuroImage (2008)

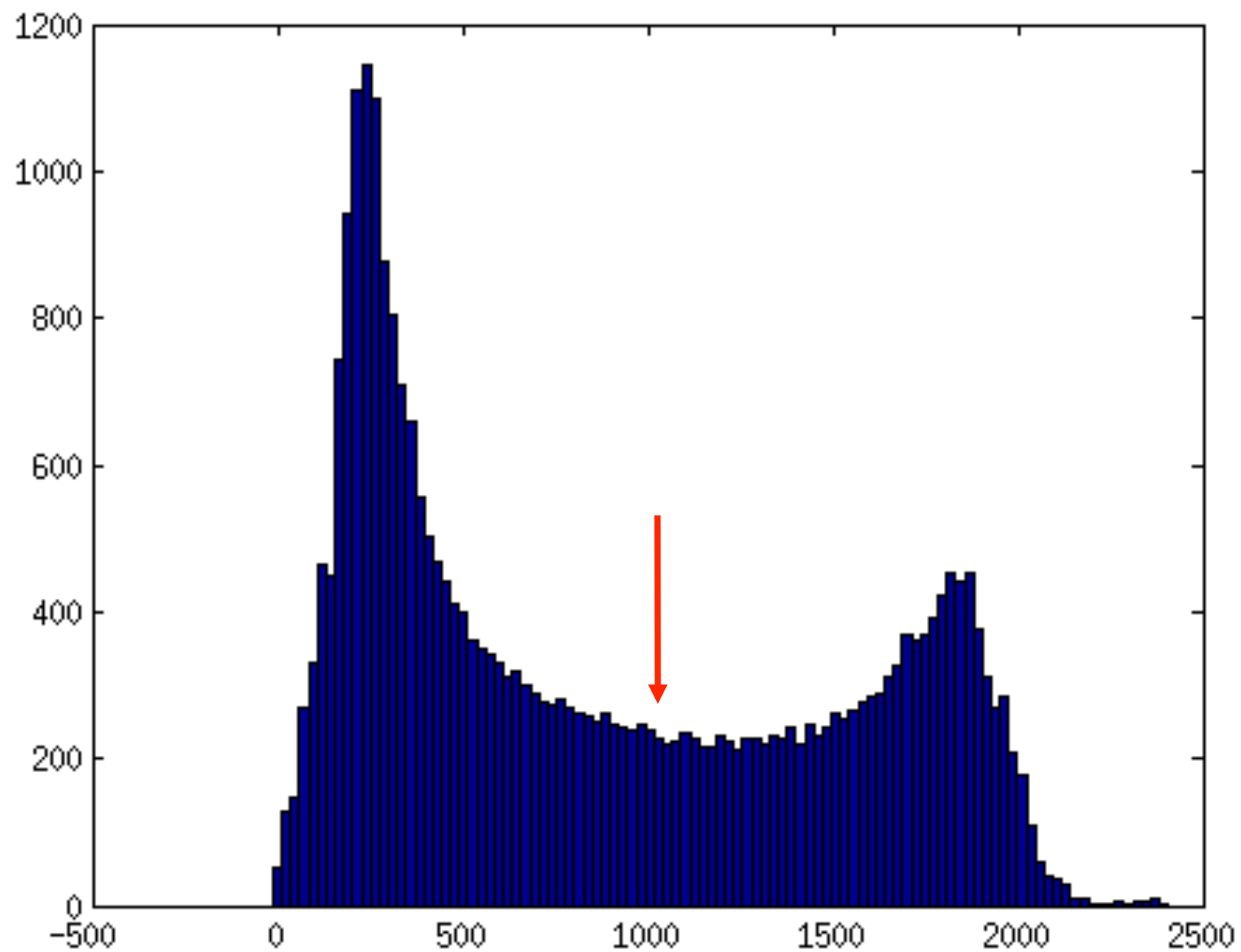
RETROICOR - RESPIRATORY PHASE



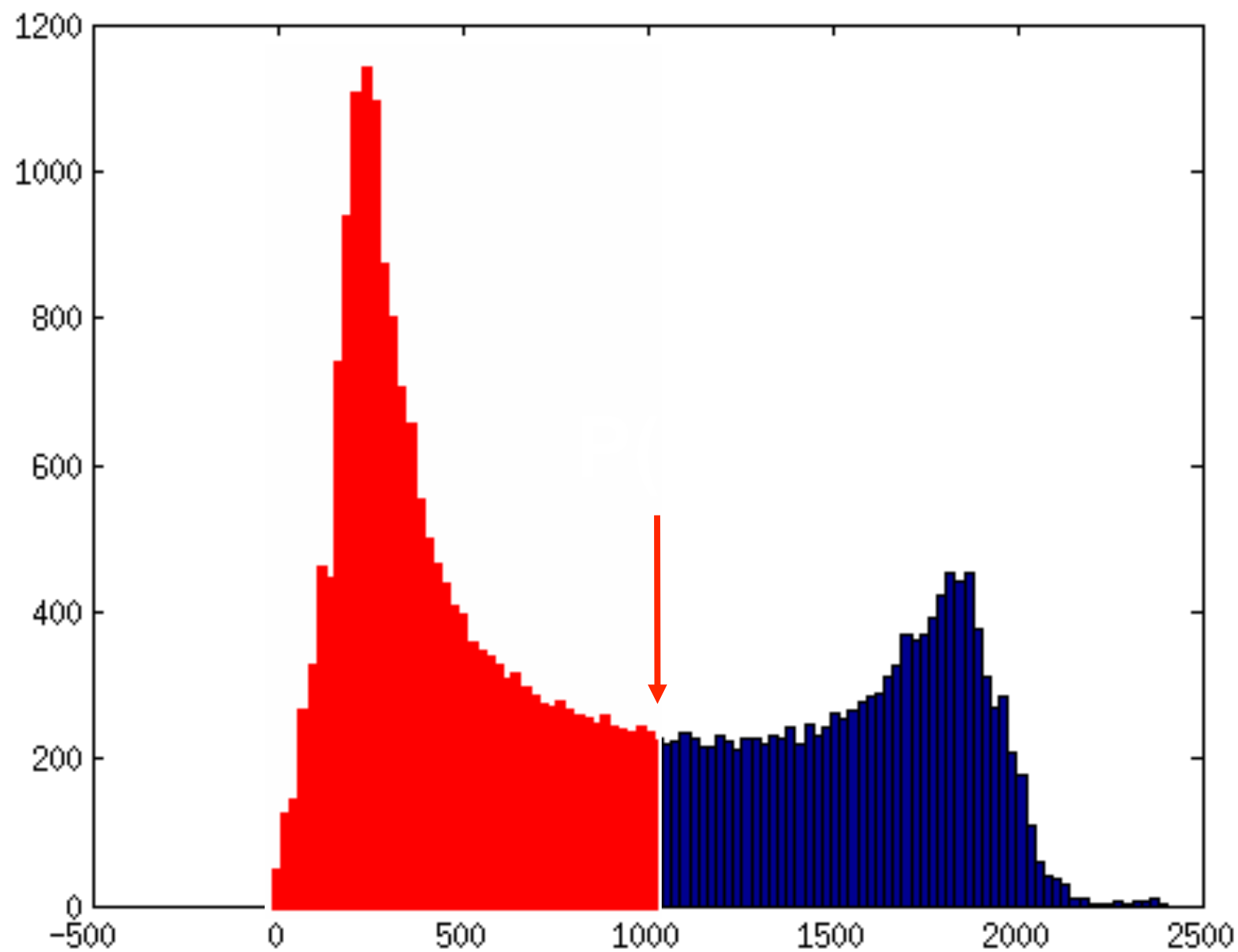
RETROICOR - RESPIRATORY PHASE



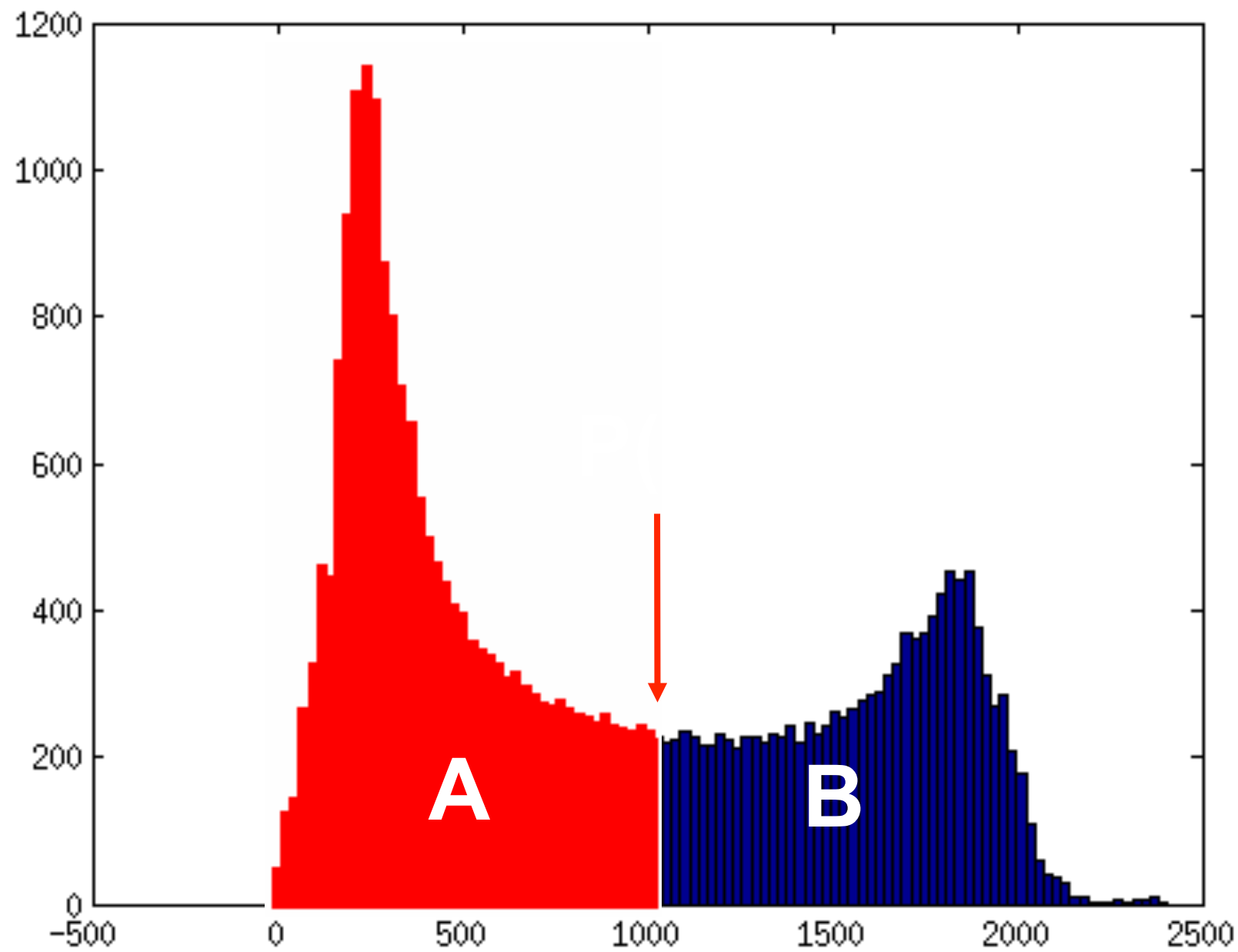
RETROICOR - RESPIRATORY PHASE



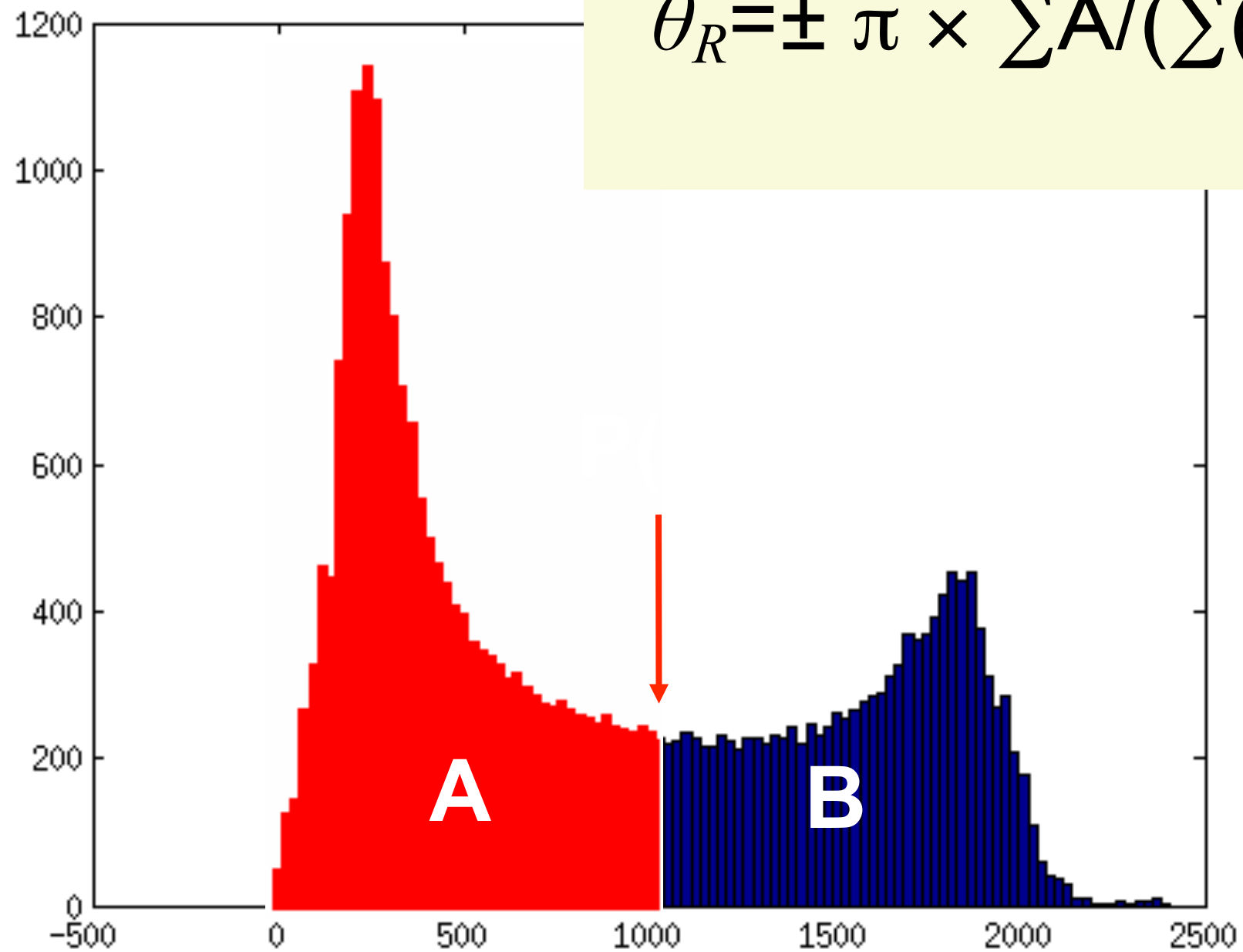
RETROICOR - RESPIRATORY PHASE



RETROICOR - RESPIRATORY PHASE



RETROICOR - RESPIRATORY PHASE



$$\theta_R = \pm \pi \times \frac{\sum A}{\sum (A+B)}$$

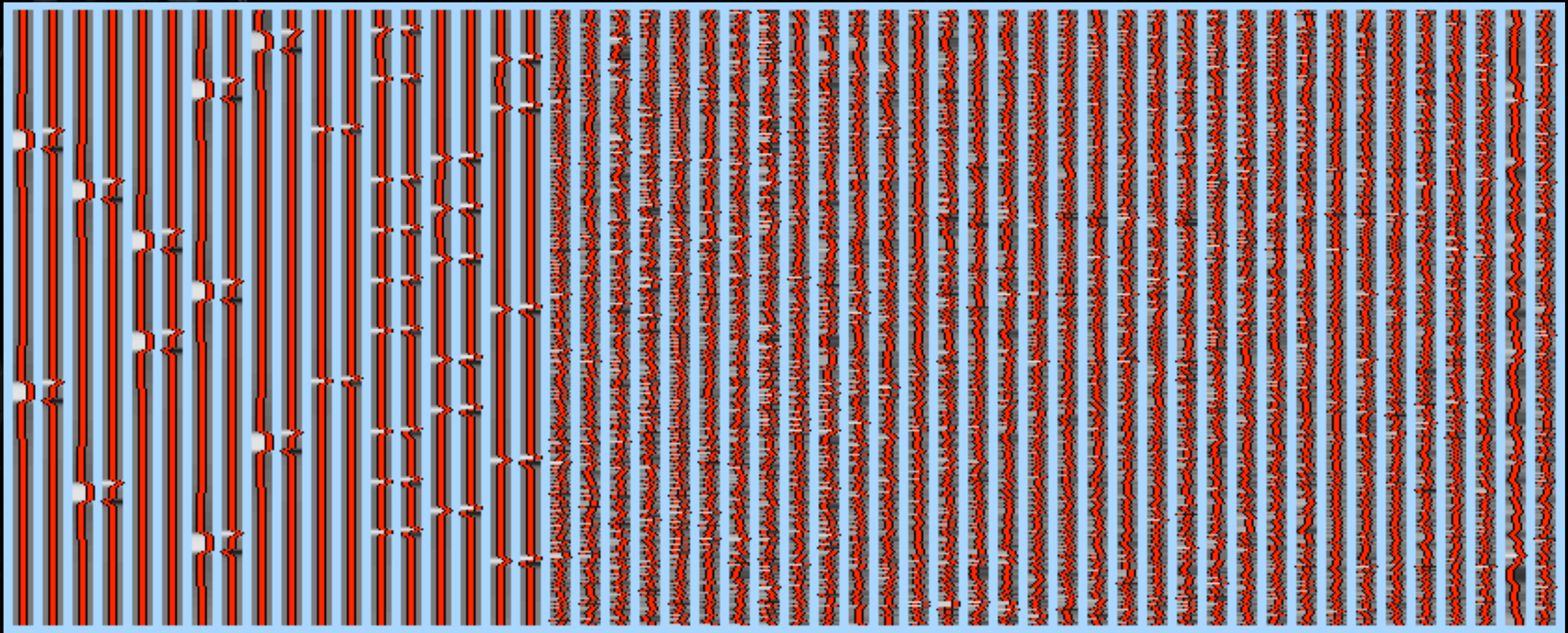
RETROICOR

- Fourier analysis (RETROICOR) can be used to model noise (*Glover et al. MRM, 2000*)
- Use 4 terms for cardiac (sine/cosine)
- Use 4 terms for respiratory (sine/cosine)
- Use 2 terms for interaction:

$$\sin/\cos(\alpha\theta_C \pm \beta\theta_R) \quad (\alpha, \beta = 1, 2)$$

- These are fed into Feat along with the experimental design and should (hopefully) explain most of the physiological noise in the images

Model





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- Spinal fMRI

A sagittal MRI scan of the spine, showing the vertebrae and intervertebral discs in grayscale. The spine curves from the top left towards the bottom right of the frame.

What's the POINT??!!

- Reduced number of subjects required to detect group effect
- Possibility to draw conclusions from $N=1$ expts?
- Greater accuracy of parameter estimates
- Ability to extract meaningful signal from difficult regions e.g. brainstem, spinal cord, areas near Circle of Willis: VTA, hippocampus, amygdala

Increased TSNR

TSNR

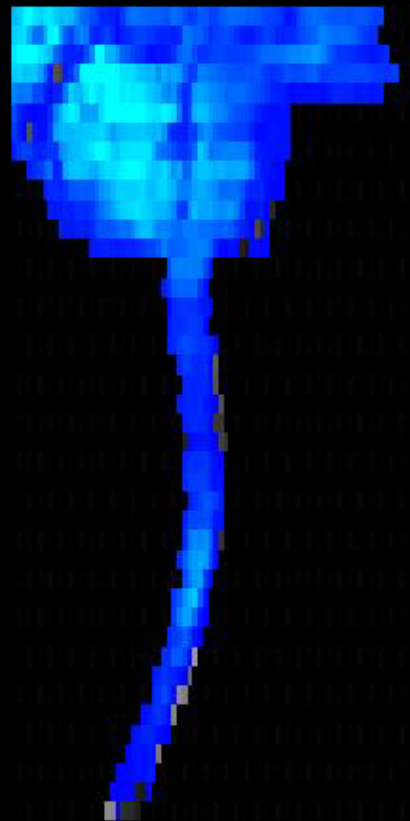
raw

100

PNM

difference (%)

100



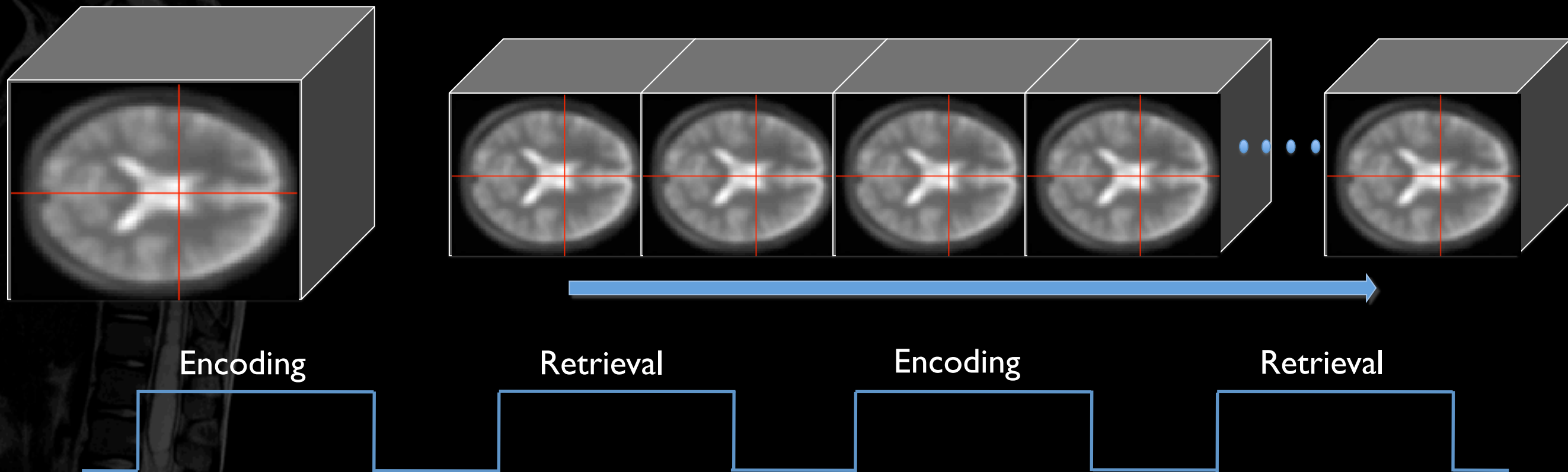
10



10

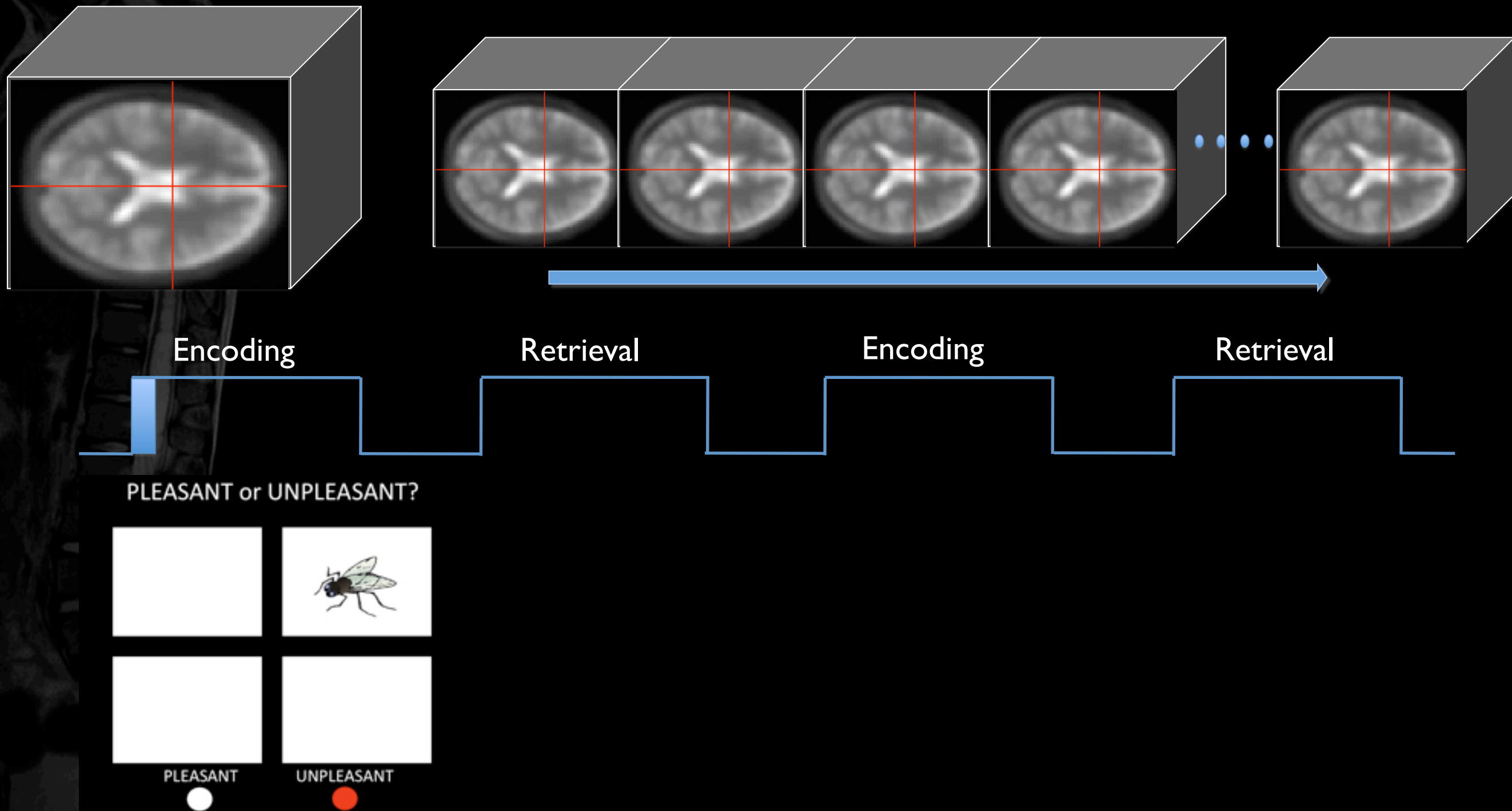
The fMRI-adapted version of the Placing Test

During the fMRI experiment subjects are asked to perform a visuo-spatial pair associates learning test ("fMRI adapted" Placing Test)



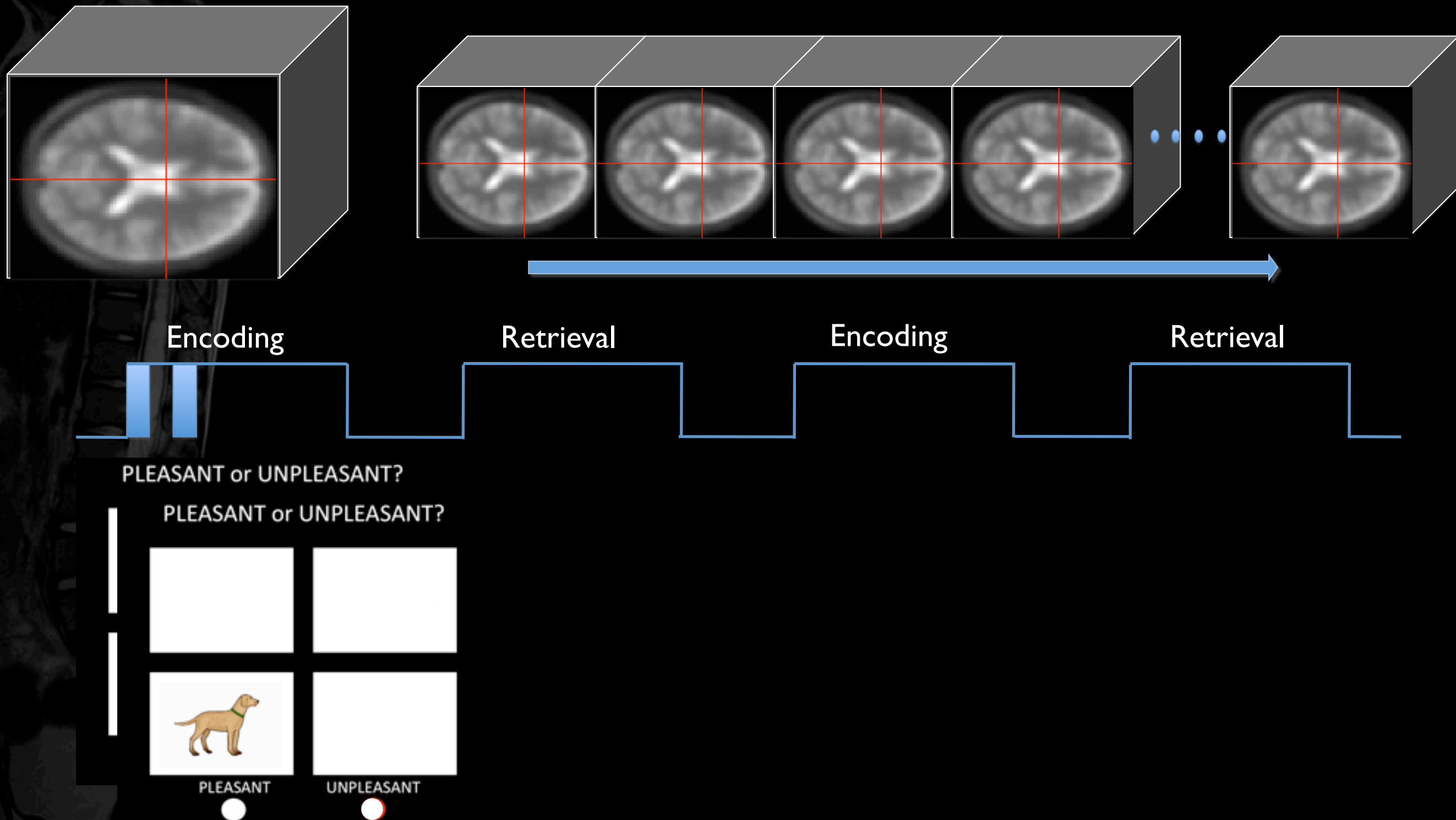
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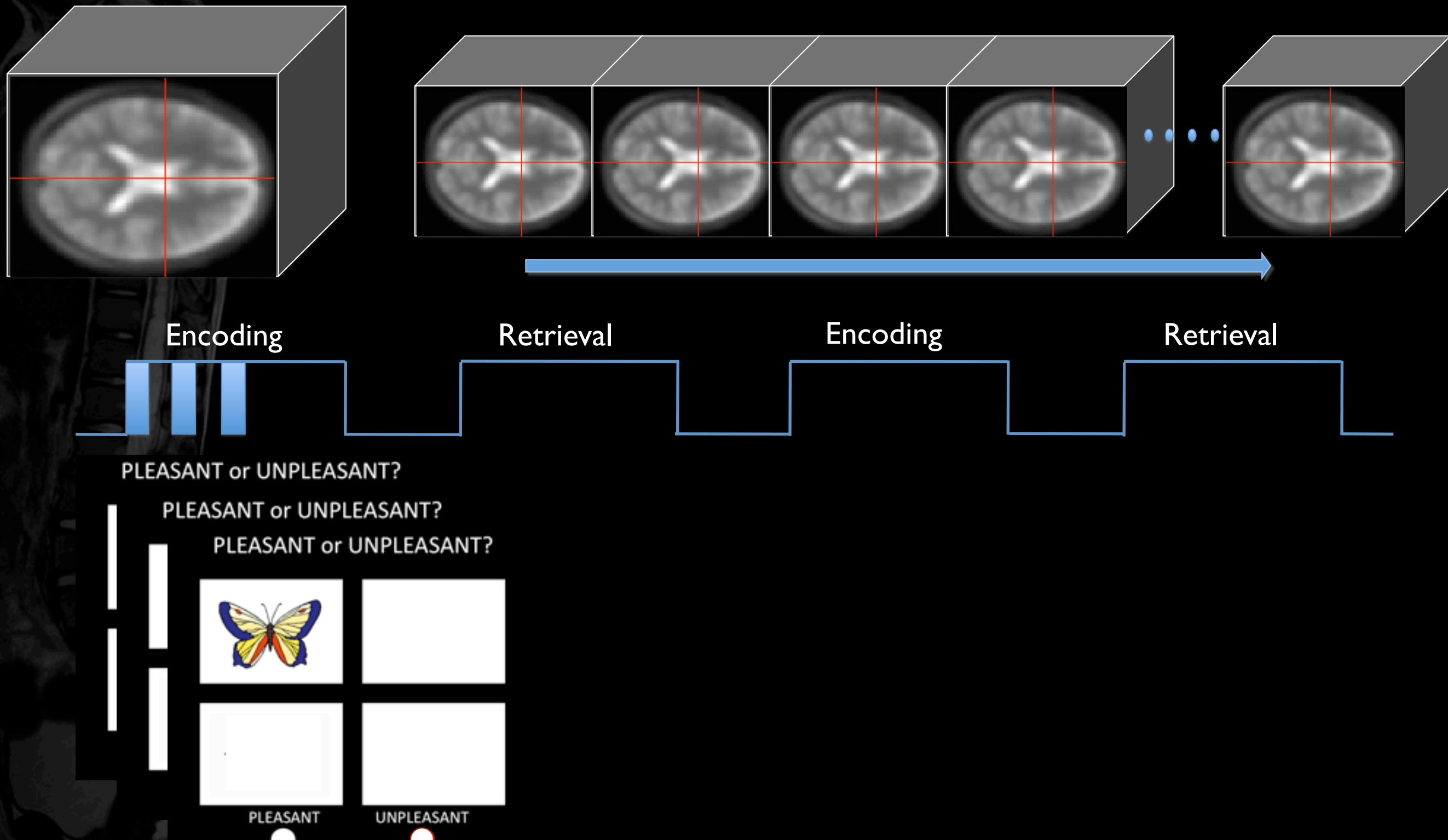
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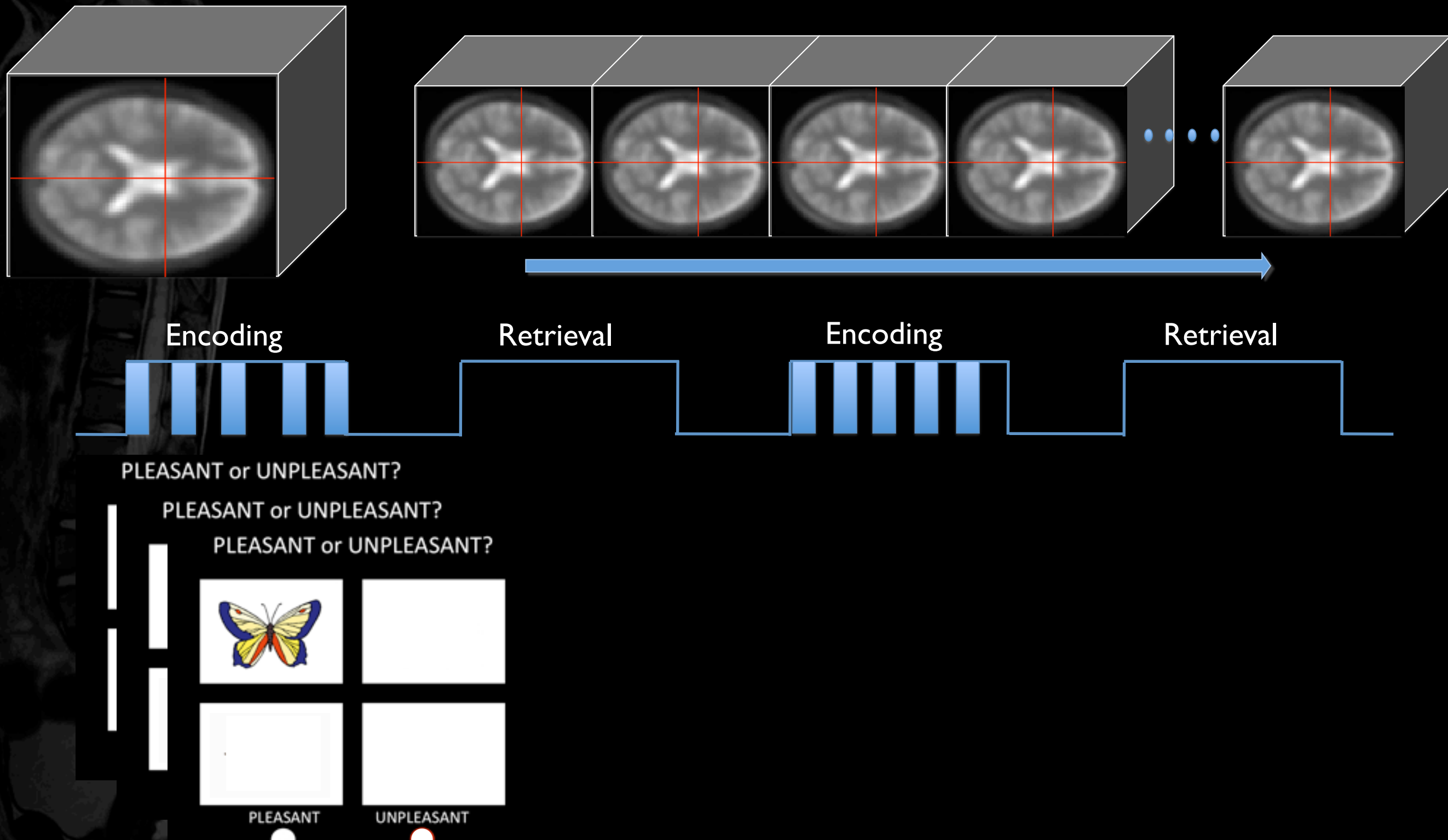
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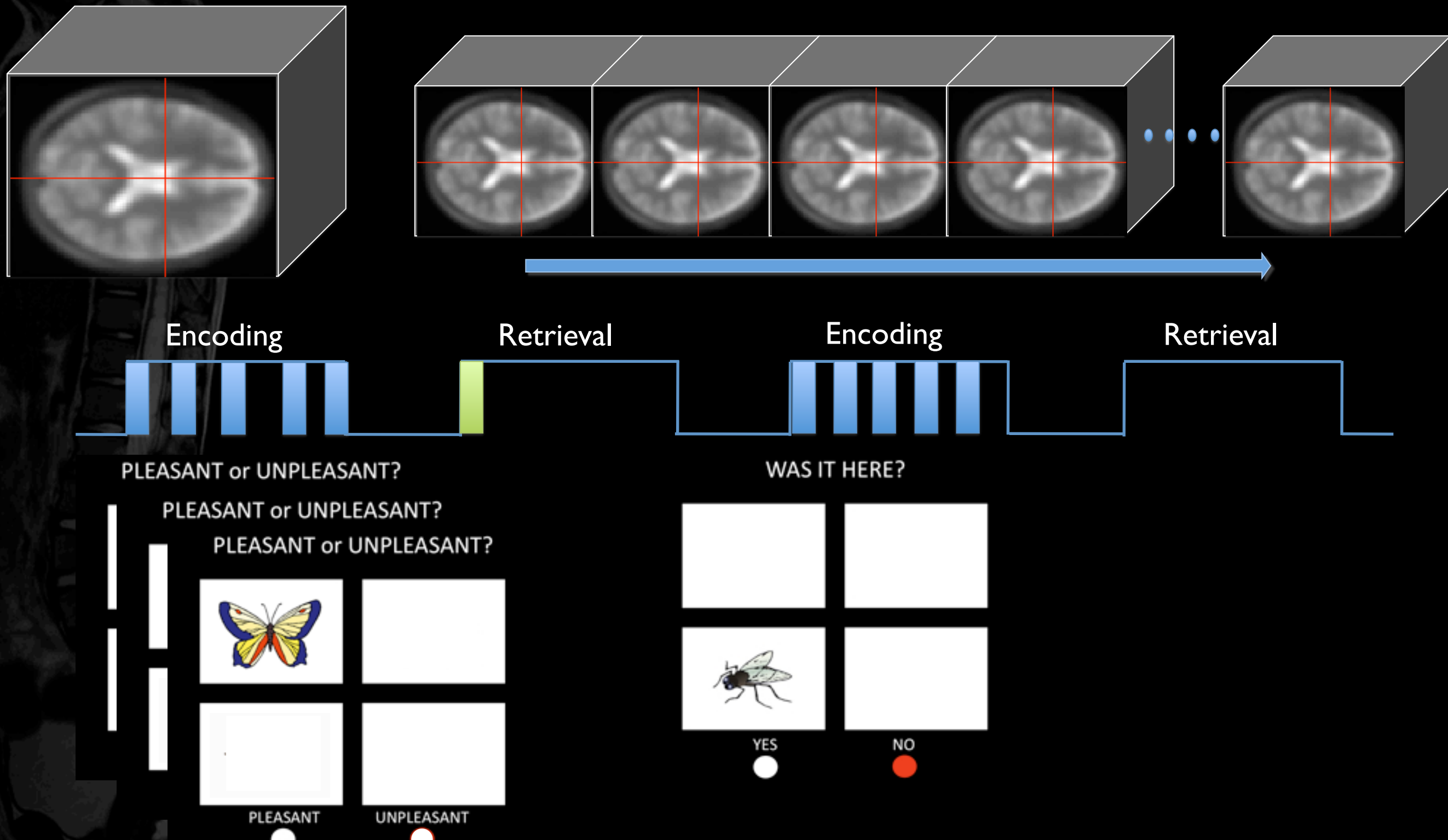
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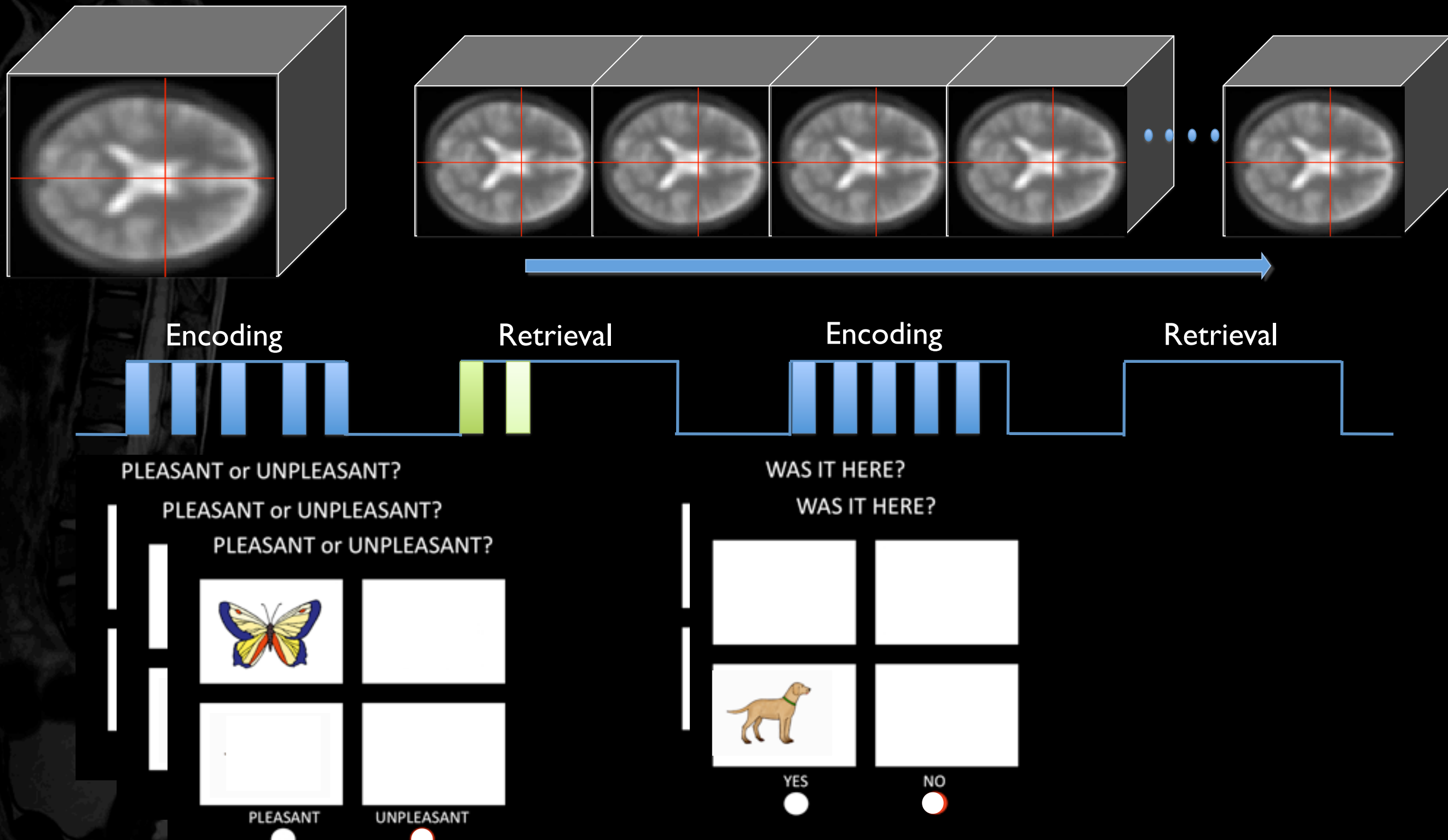
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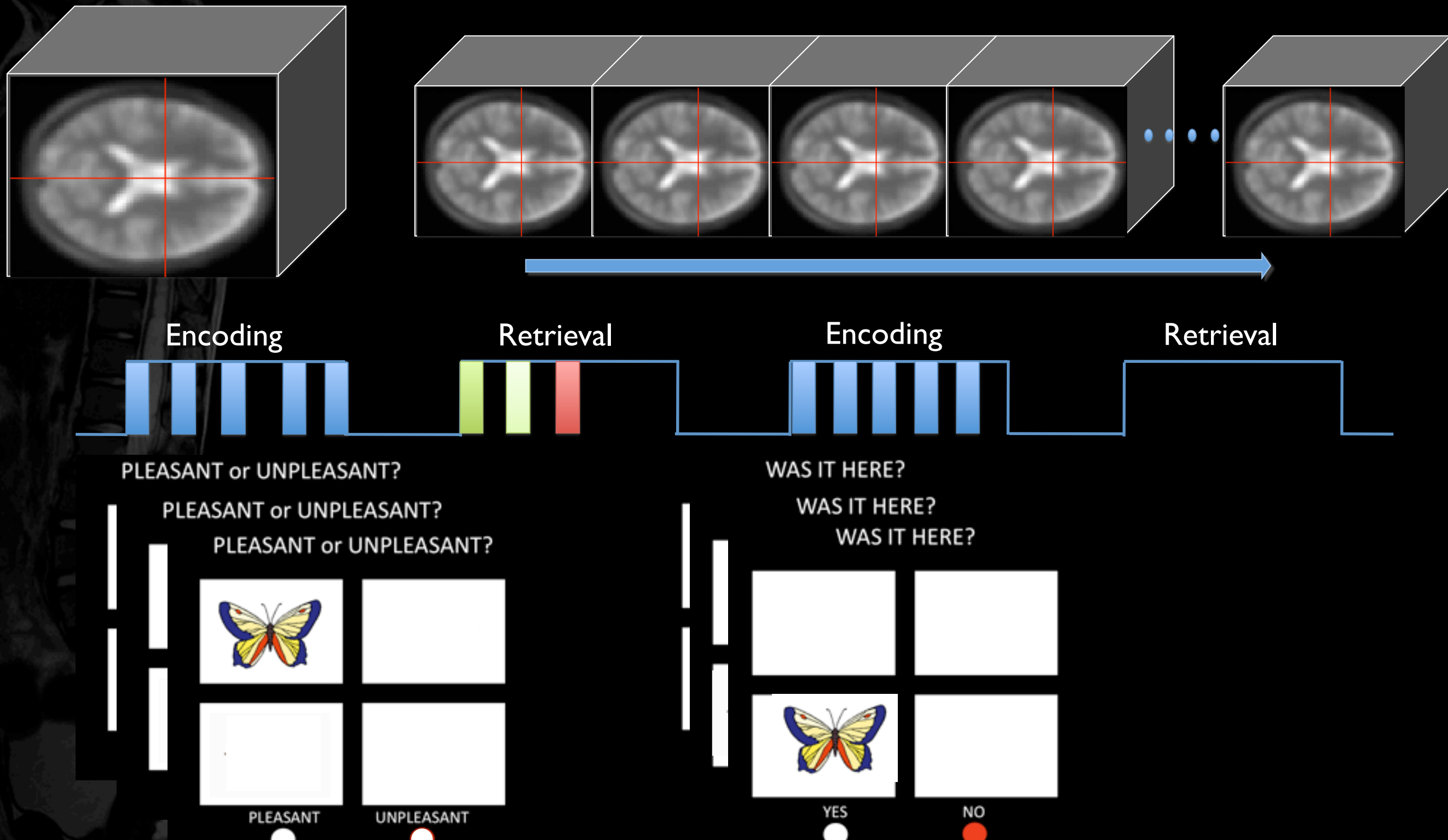
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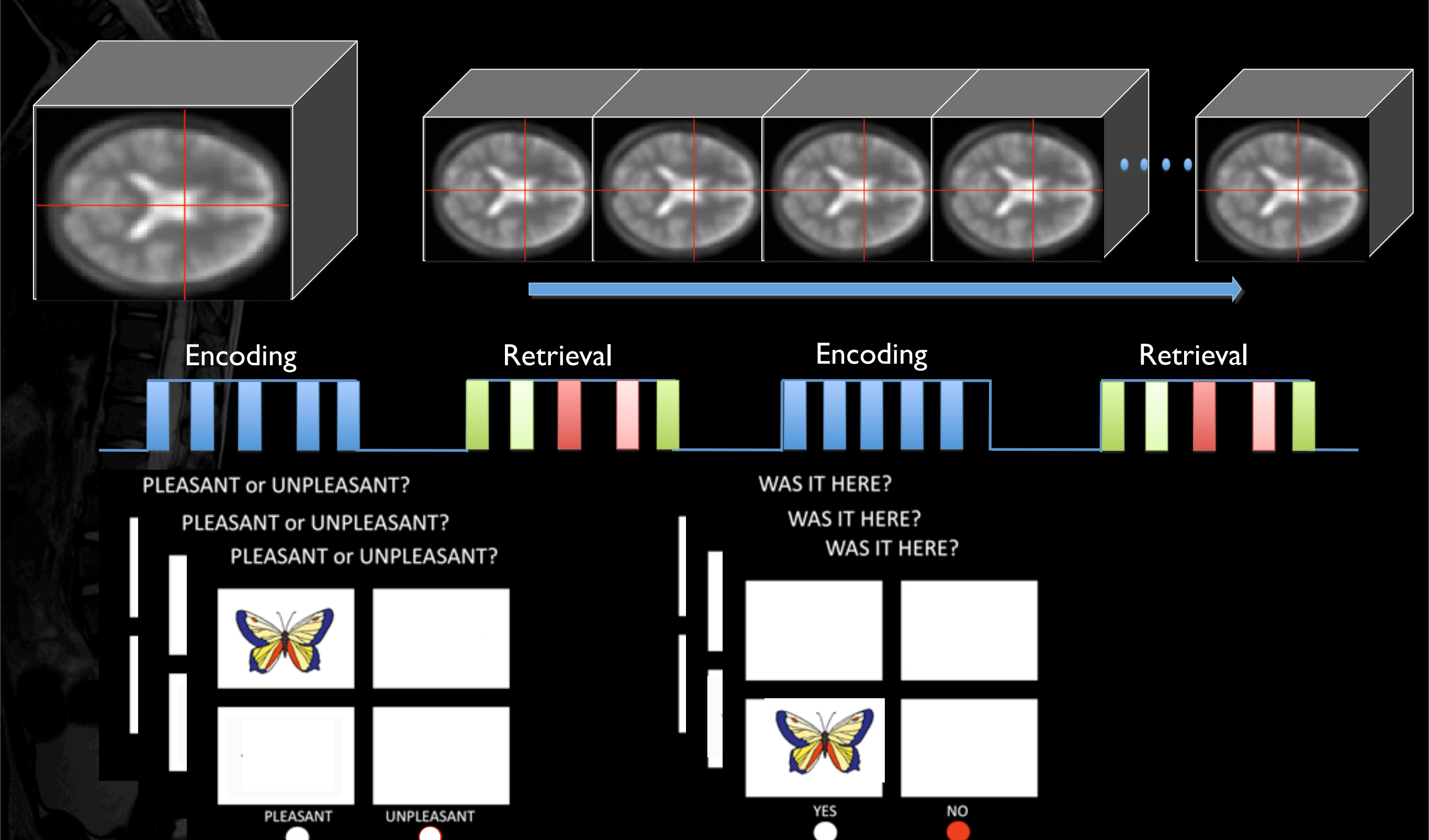
The fMRI-adapted version of the Placing Test

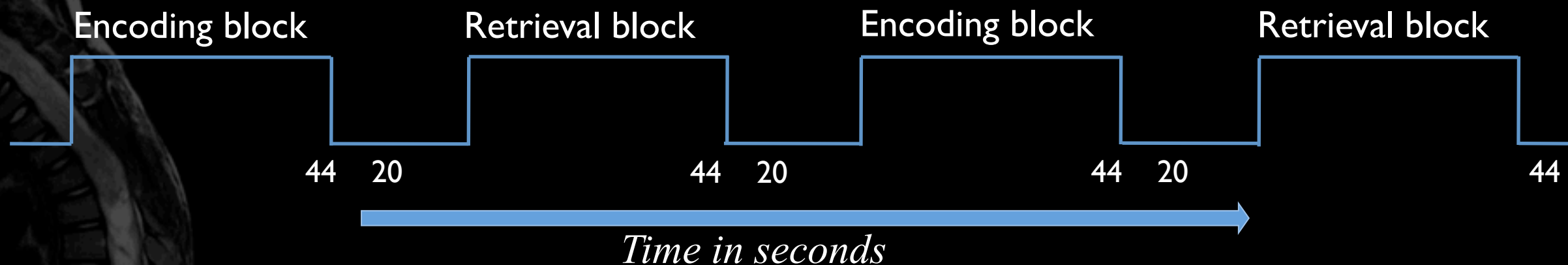
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The fMRI-adapted version of the Placing Test

*During the fMRI experiment subjects are asked to perform a visuo-spatial pair associates learning test
("fMRI adapted" Placing Test)*





Encoding block

PLEASANT or UNPLEASANT?

PLEASANT or UNPLEASANT?

PLEASANT or UNPLEASANT?

PLEASANT or UNPLEASANT?

PLEASANT or UNPLEASANT?



PLEASANT



UNPLEASANT



Retrieval block

WAS IT HERE?

WAS IT HERE?

WAS IT HERE?

WAS IT HERE?

WAS IT HERE?



YES



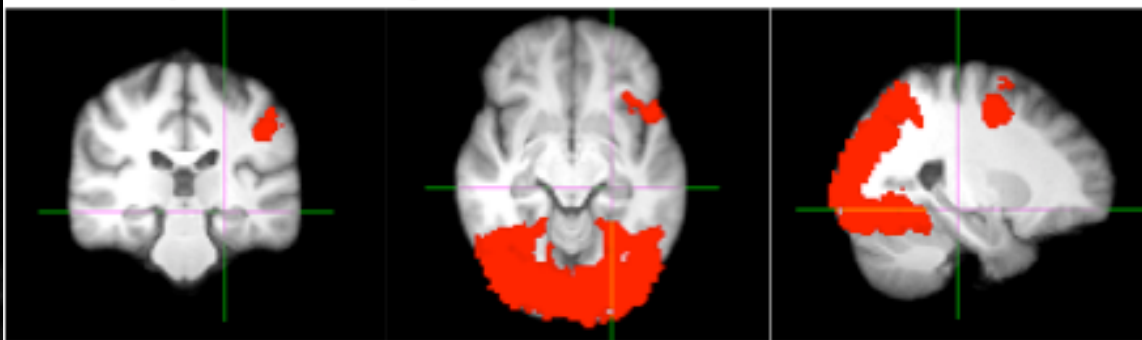
NO



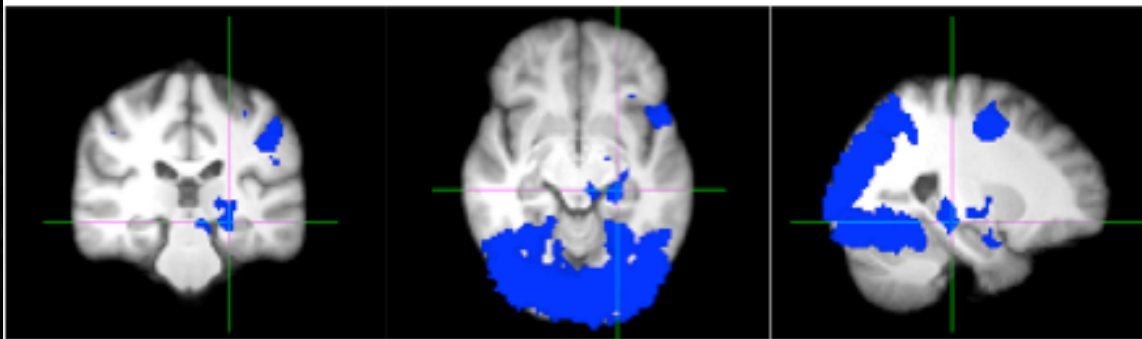
5
associates
per block
6 secs
each

Hippocampal activity

Non-Physio Encoding (Thresh 2.0)

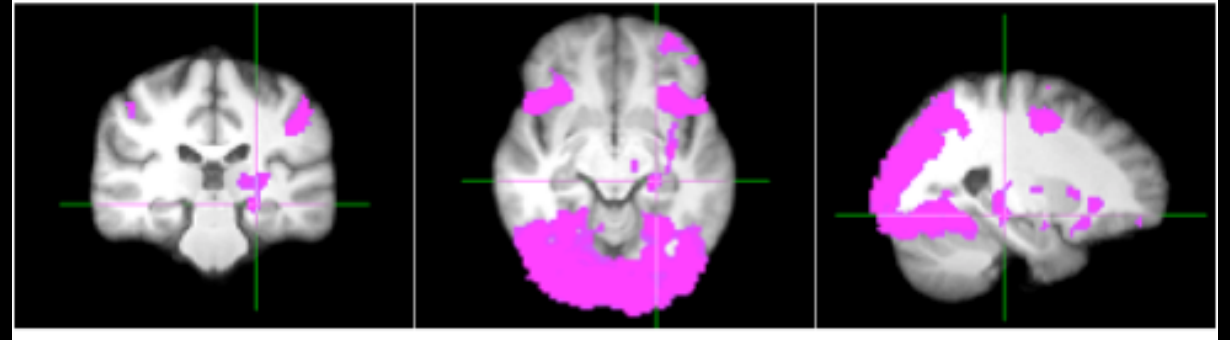


Physio Encoding (Thresh 2.0)

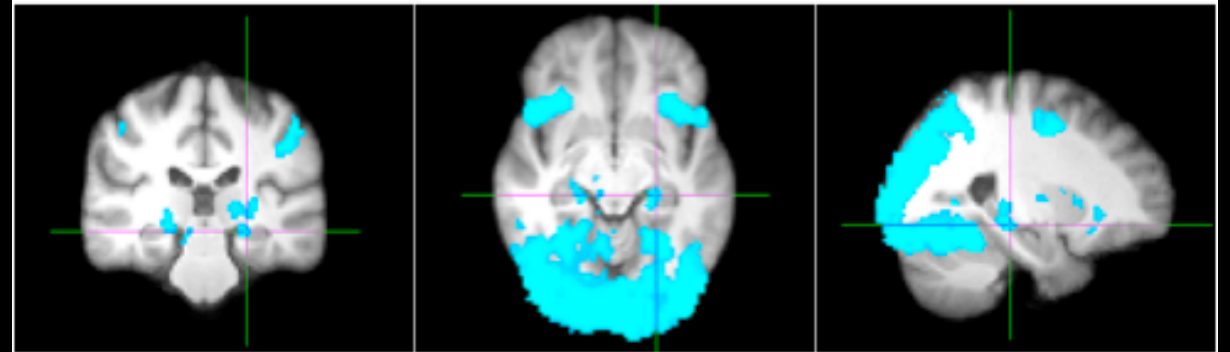


encoding, cluster
corrected $p < 0.05$

Non-Physio Retrieval (Thresh 2.3)



Physio Retrieval (Thresh 2.3)



retrieval, cluster corrected
 $p < 0.05$

Brainstem experiments

METHODS

Subjects: To date, six right-handed healthy volunteers (3 male, 3 female; age: 26 ± 2.17).

Physiological monitoring: Respiratory bellows, pulse oximeter and CO2 sampling via a BIOPAC device.

Paradigm: 2 runs of pain stimuli and 2 runs of vibrotactile stimuli, each run testing either a coronal $2 \times 2 \times 2 \text{ mm}^3$ or an axial $1.5 \times 1.5 \times 3 \text{ mm}^3$ resolution acquisition.

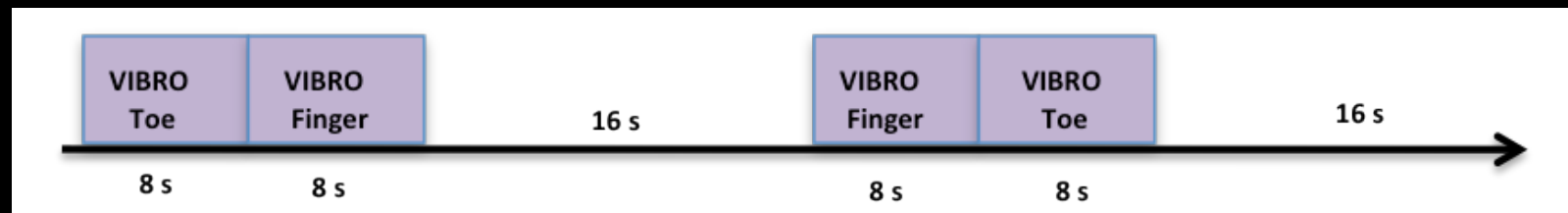


Figure 1A. Vibrotactile paradigm- subjects received 20 blocks of vibrotactile stimuli to the right index finger and the right hallux delivered pseudorandomly at 30Hz via a Piezo-electric vibrotactile device.

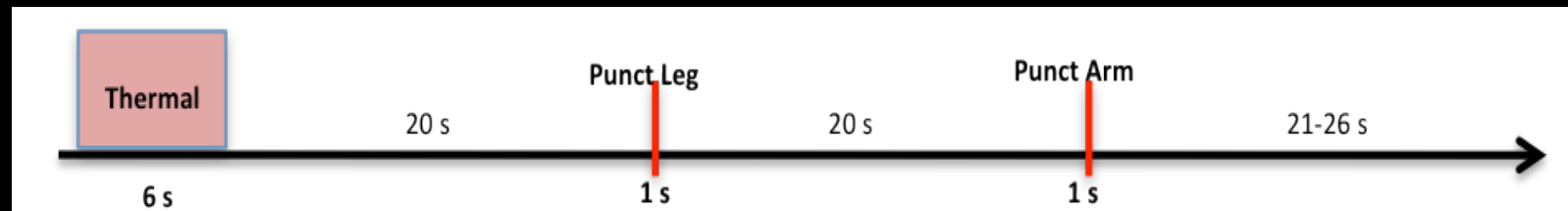
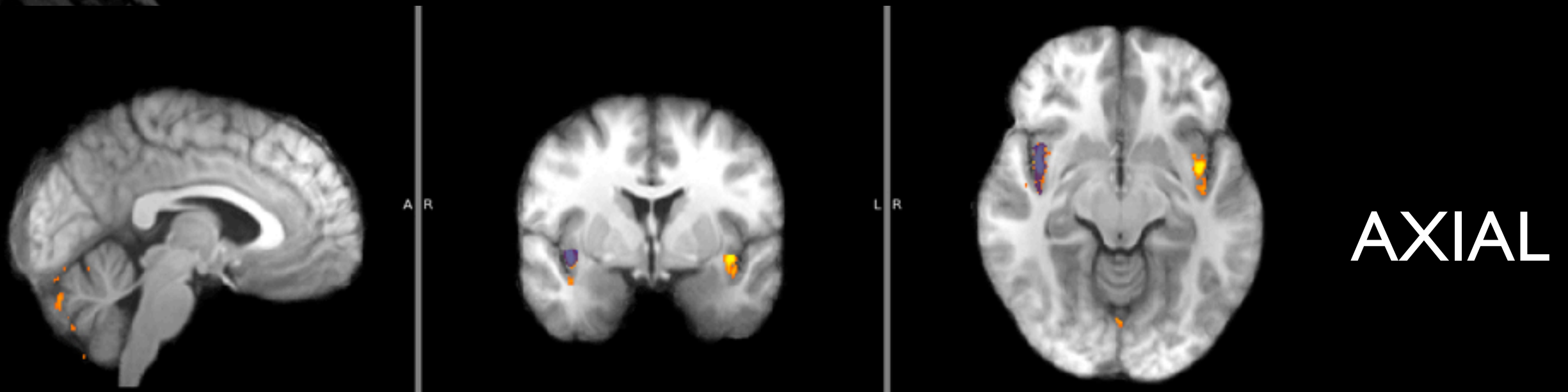


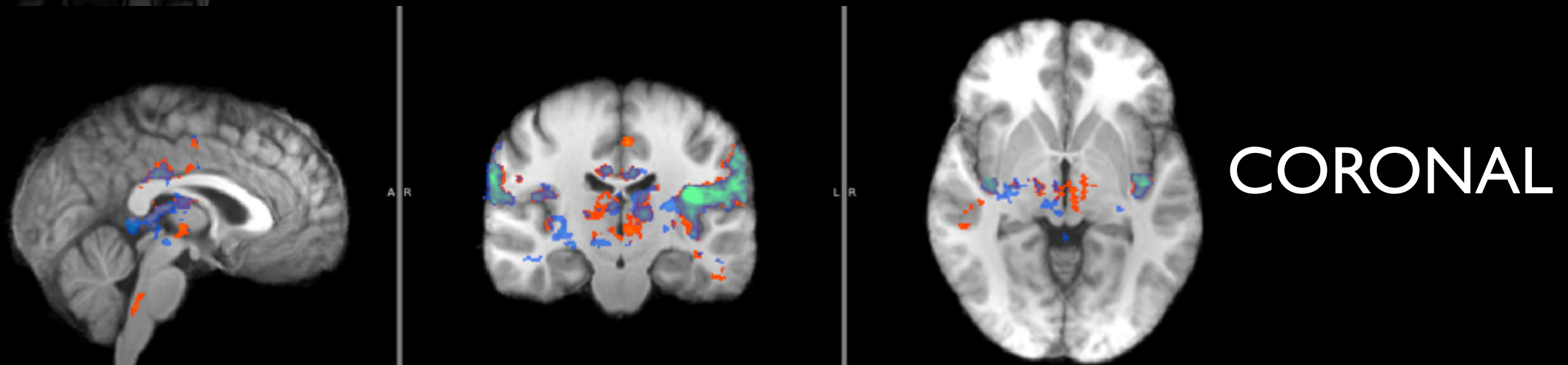
Figure 1B. Pain paradigm- subjects received 15 thermal stimuli to the right volar forearm, delivered with a MEDOC Pathway CHEPS device and thresholded at 6/10 on an 11-point pain rating scale. The thermal stimuli were separated by two punctate stimuli delivered pseudorandomly between the right arm and right leg using a 512mN punctate probe.

Analysis: Data was processed using FMRIB software library (FSL) tools. Physiological data was processed in MATLAB.

Pain- thermal



N=6, Group mean (Fixed effects) $Z=1.8$ $p<0.05$



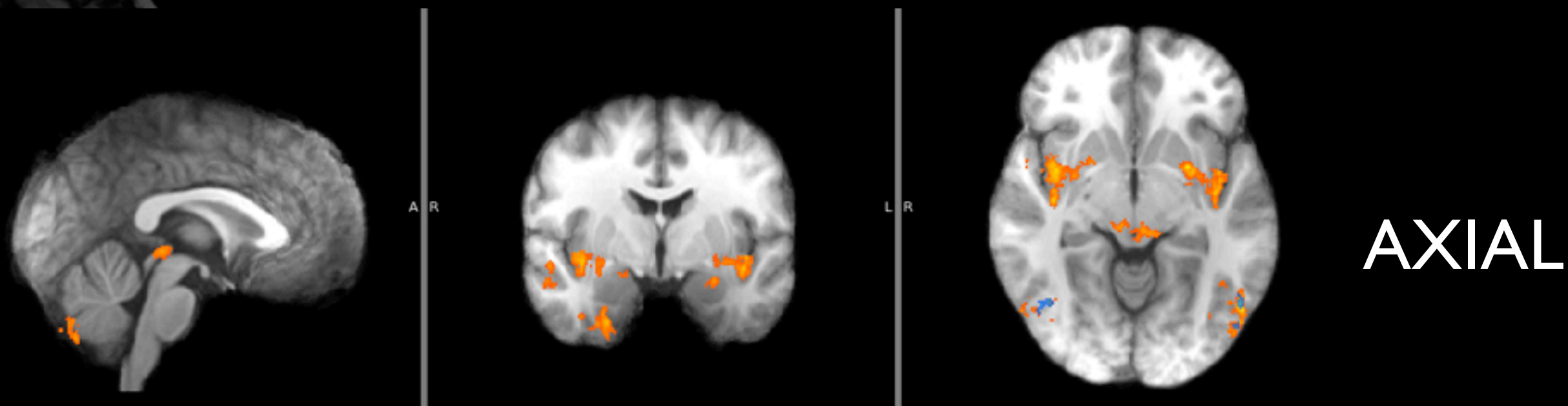
With PNM



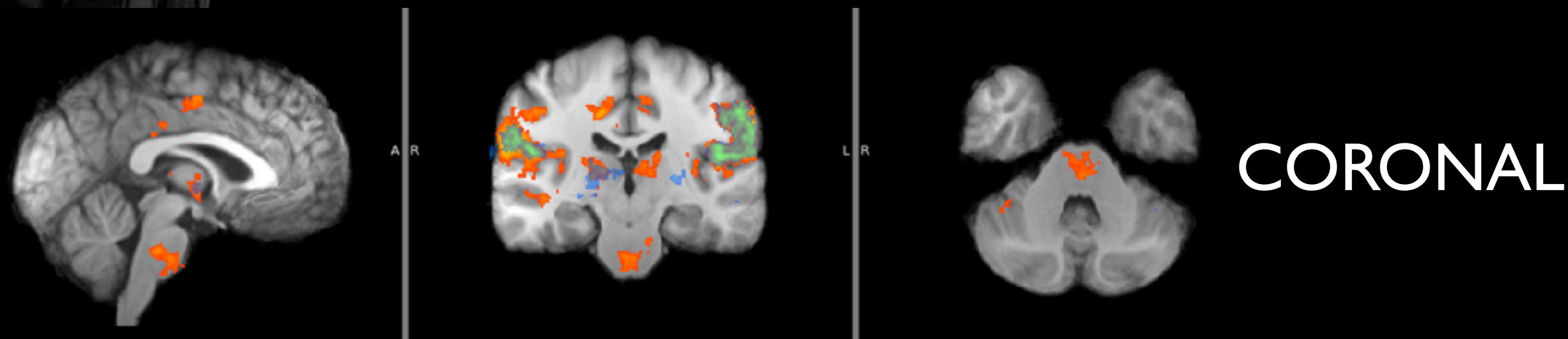
Without PNM



Pain- punctate arm



N=6, Group mean (Fixed effects), $Z=1.8$ $p<0.05$



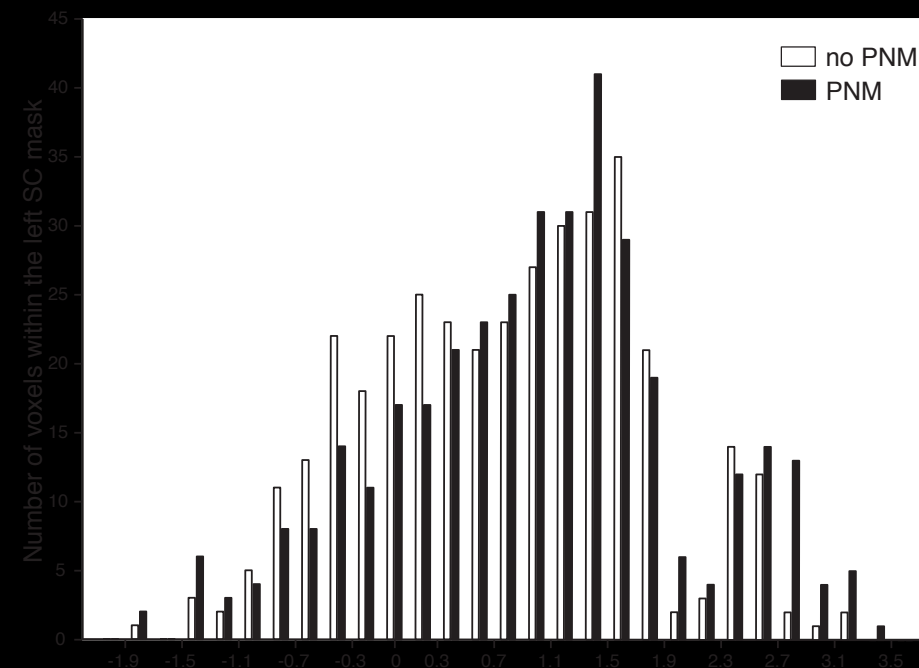
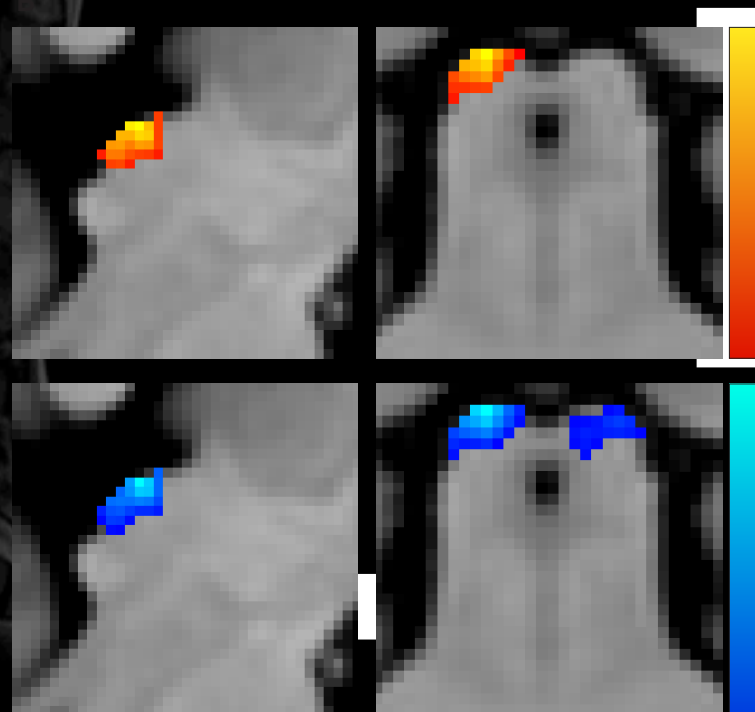
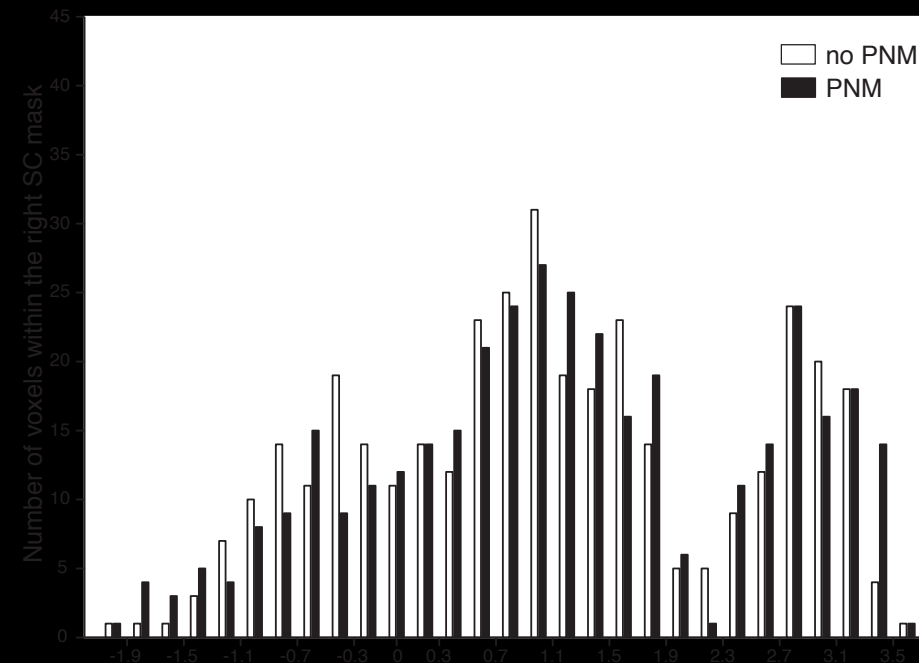
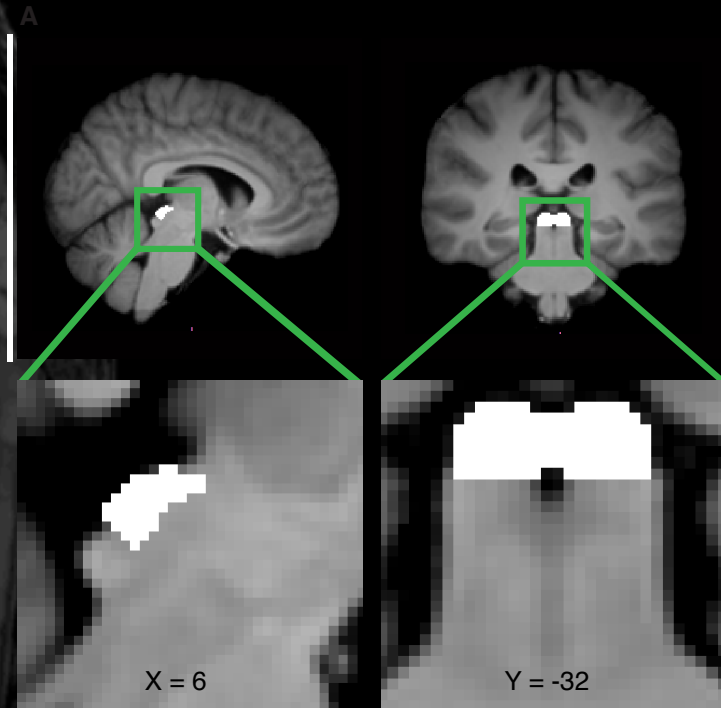
With PNM ■

Without PNM ■

Visual paradigm

- 10 healthy controls
- Coronal oblique acquisition through brainstem (superior colliculi)
- Philips 3T scanner, ECG and respiratory bellows
- Smoothly rotating semi circle made of alternating black and white checks that scaled linearly with eccentricity
- Checks reversed contrast at 8Hz and the semi-circle rotated at 1Hz. Each presentation lasted for two seconds, random ITI, jittered
- Data analysis MATLAB script, and 4D regressors in FSL

Visual paradigm



Limbrick-Oldfield et al (2012). NeuroImage 59:1230-1238.

A sagittal MRI scan of a human spine, showing the vertebrae and intervertebral discs. The image is in grayscale and occupies the left side of the slide.

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- Is it worth the hassle?
- **Spinal FMRI**



Spinal fMRI

- Why? Understanding processes occurring at the spinal level (e.g. sensorimotor)
- Pain - central sensitisation, also brainstem
- Greater influence of physiological noise as you move down the neuraxis:

brain << (brainstem < spinal cord)



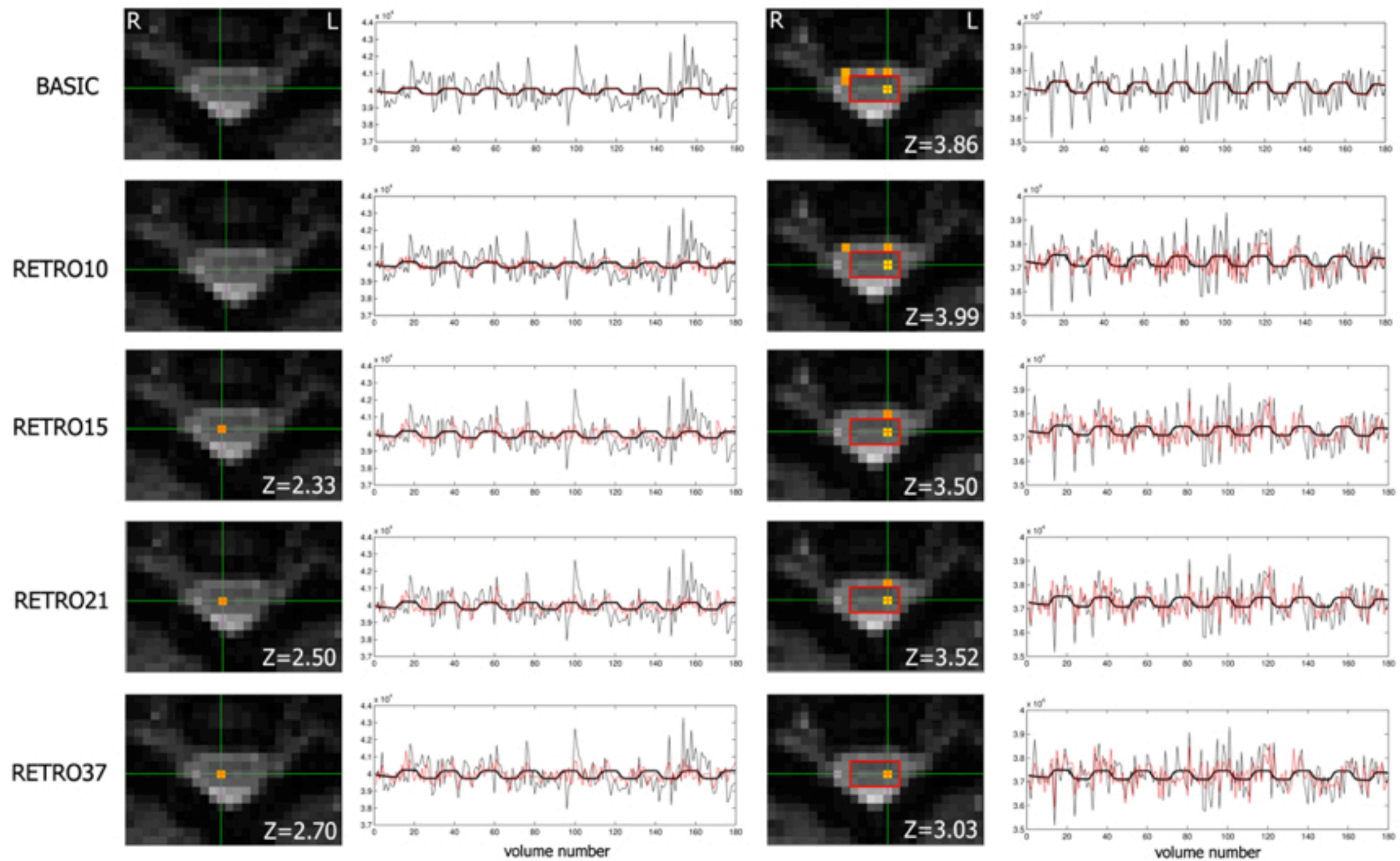
Physiological Noise Model (PNM)


- Uses a sum of sine and cosine terms
- Empirically defined regressor (CSF)
- Modelled using the GLM in Feat
- Available in FSL5

Brooks et al (2008) Physiological noise modelling for spinal functional magnetic resonance imaging studies.
NeuroImage 39:680-692.

RIGHT HAND STIMULATION

LEFT HAND STIMULATION





Useful info

- <http://fsl.fmrib.ox.ac.uk/fsl/fslwiki/PNM>
- <http://www.fmrib.ox.ac.uk/Members/jon/physiological-noise-correction>

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Acknowledgments

- Mark Jenkinson, Matthew Webster, Karla Miller, Jesper Anderson, Christian Beckmann, Irene Tracey (FMRIB)
- Yazhuo Kong (FMRIB)
- Eve Limbrick-Oldfield (Imperial College)