



# From the Basics of MRI and PET to Hybrid MR-PET

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GERMANY

# On the Road Towards Metabolic Imaging: Recent Advances

Shah (2014) *Brain Structure and Function* (in press)

# Quantitative Imaging

N. J. Shah et al.

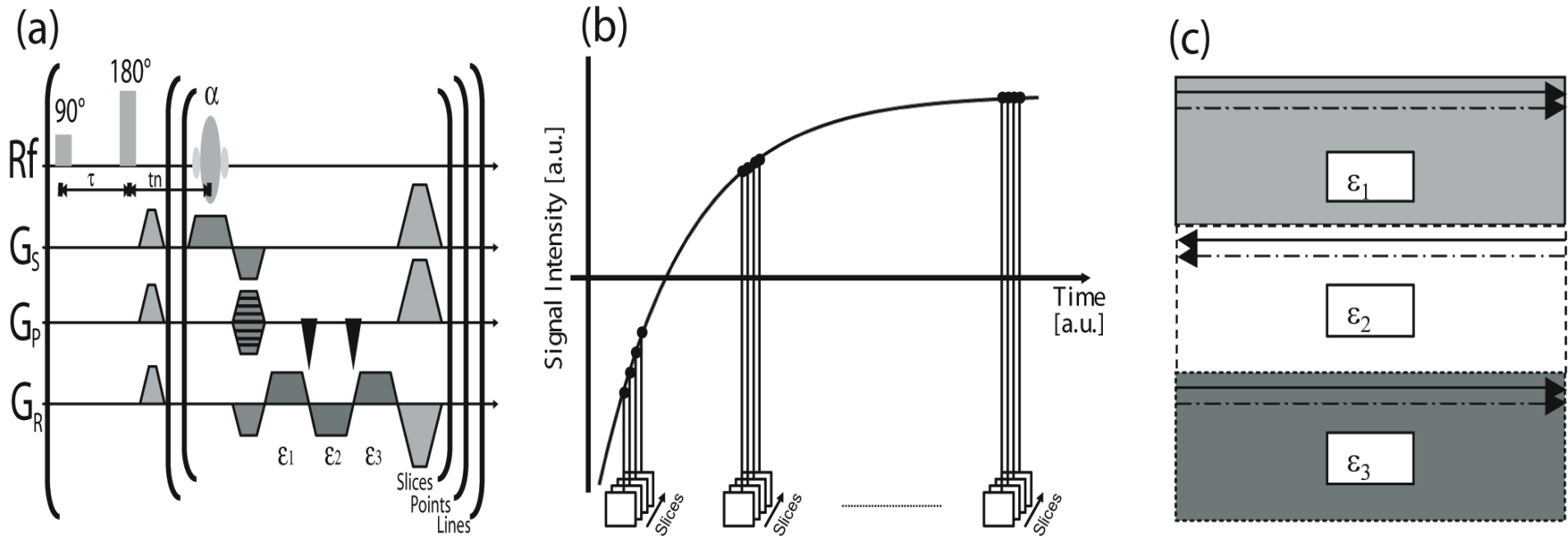
# T<sub>1</sub> Mapping

Look and Locker, Rev. Sci. Instr. 41: 250-251

Deichmann and Haase, JMR 1992 96: 608-612

Deichmann et al., MRM 1999 42: 206-209

# Implementation



## TAPIR ( $T_1$ mApping of Partial Inversion Recovery)

Shah et al.,; US Patent No.: 6,803,762

Shah et al., NeuroImage: 2001 14(5): 1175-85

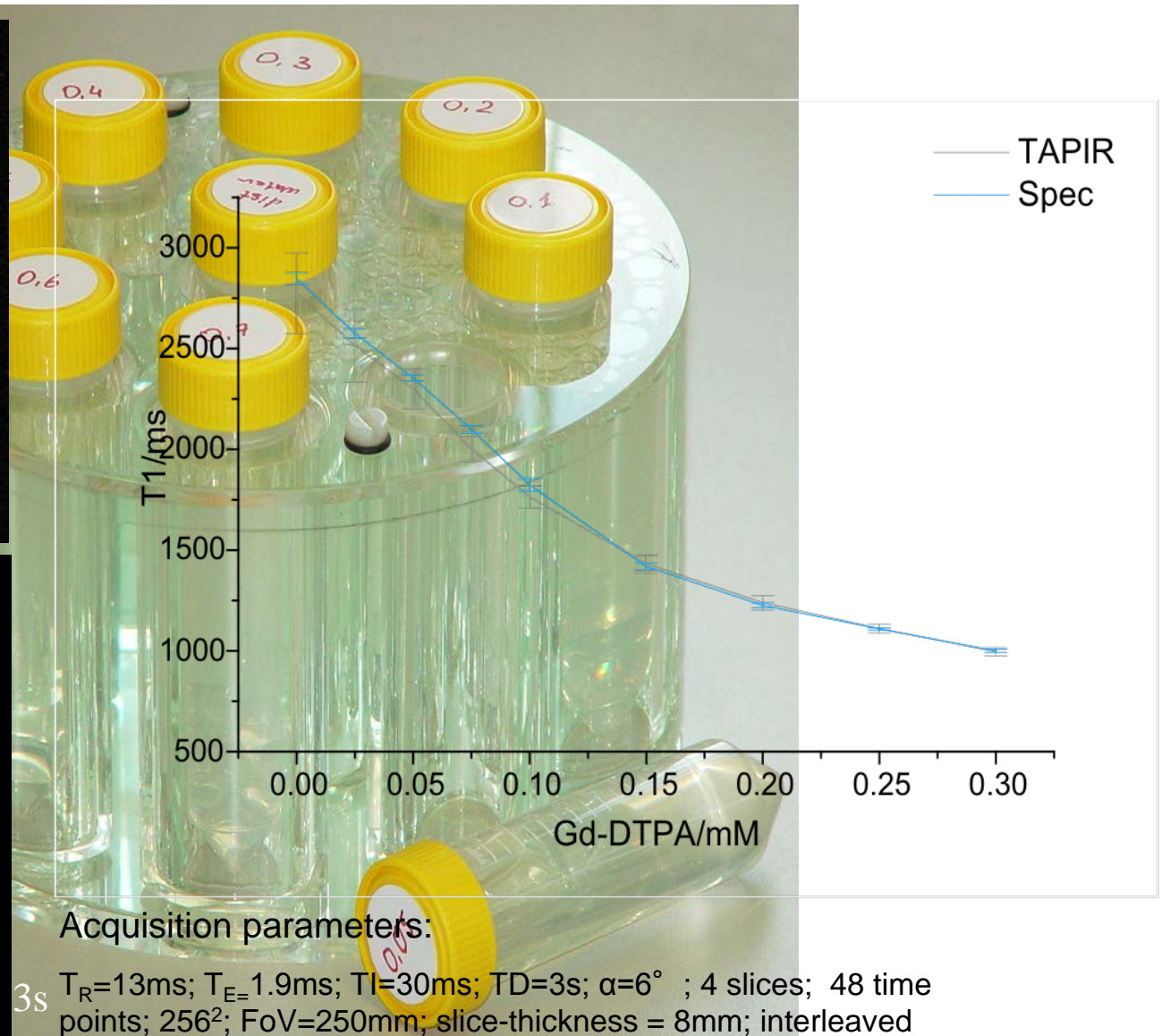
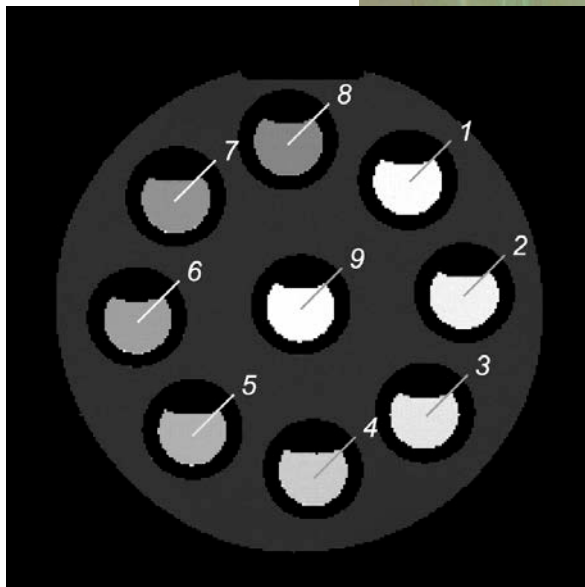
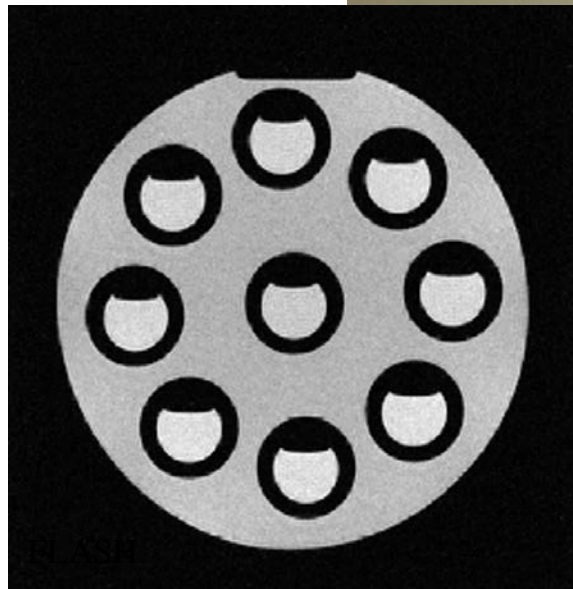
Steinhoff et al., Magn. Reson. Med.: 46(1) 131-140 2001

Zaitsev, et al; Magn. Reson. Med.: 49(1) 1121-1132 2003

Shah et al., Hepatology: 2003 38: 1219-26

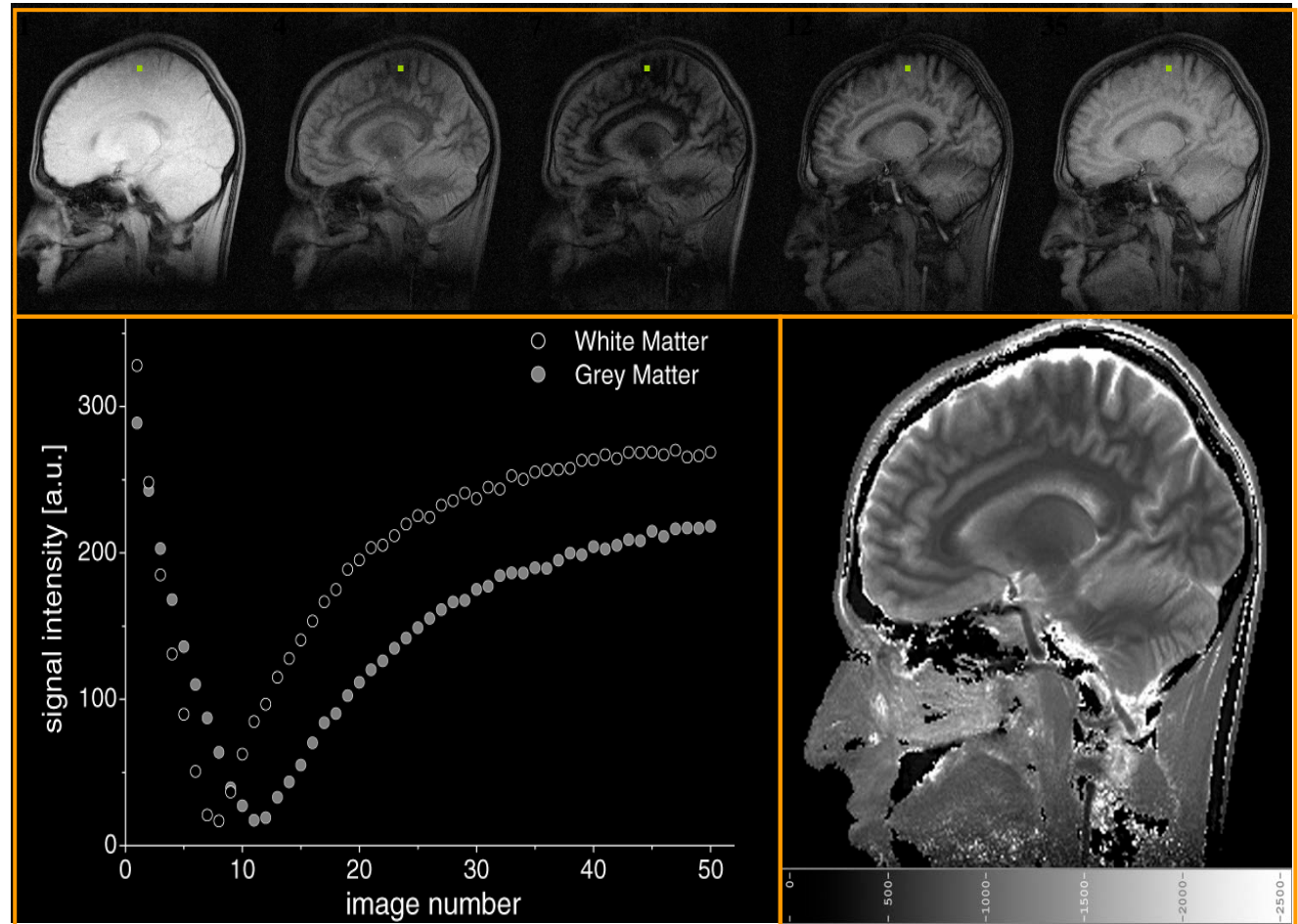
Tapir: any perissodactyl mammal of the genus Tapirus .... of South and central America and SE Asia, having an elongated snout, **three-toed** hind legs, and four-toed forelegs.

# Phantom Results



# TAPIR: *In vivo* $T_1$ Mapping

- Large number of points affords reconstruction of accurate maps
- Multi-exponential fitting is feasible
- $T_1$  mapping enables **quantitative** measurement of water content.
- $S(t) = M_0 \{1 - 2\exp(-t/T_1)\}$
- ... life is not so simple!



Shah et al.,; US Patent No.: 6,803,762

Shah et al., NeuroImage: 2001 14(5): 1175-85

Steinhoff et al., Magn. Reson. Med.: 46(1) 131-140 2001

Zaitsev, et al; Magn. Reson. Med.: 49(1) 1121-1132 2003

Shah et al., Hepatology: 2003 38: 1219-26

31 October 2015

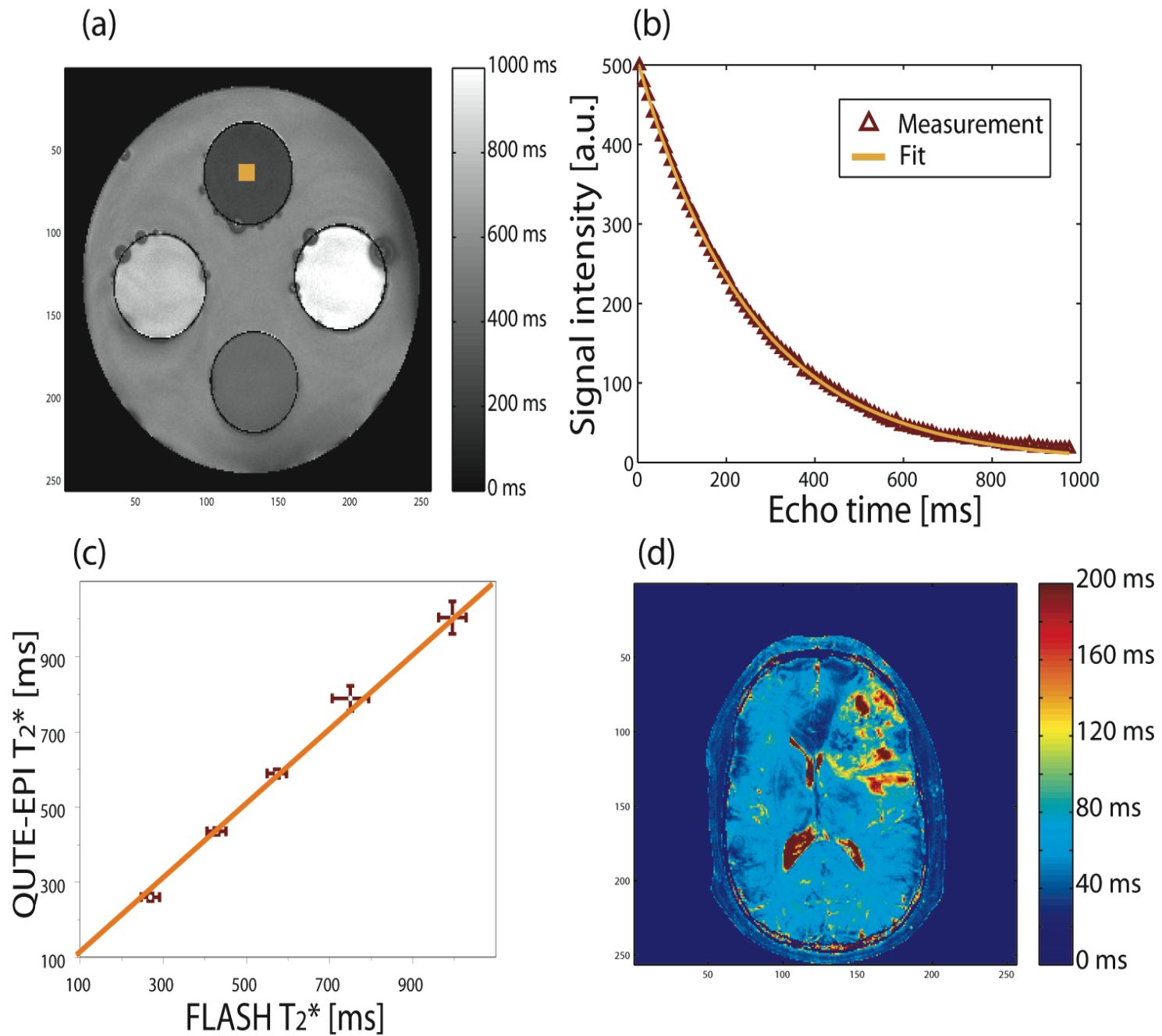
Institute of Neuroscience and Medicine

# $T_2^*$ Mapping

Mansfield: 1984 Spectroscopic Imaging (EPSI)



# Phantom and *in vivo* Results

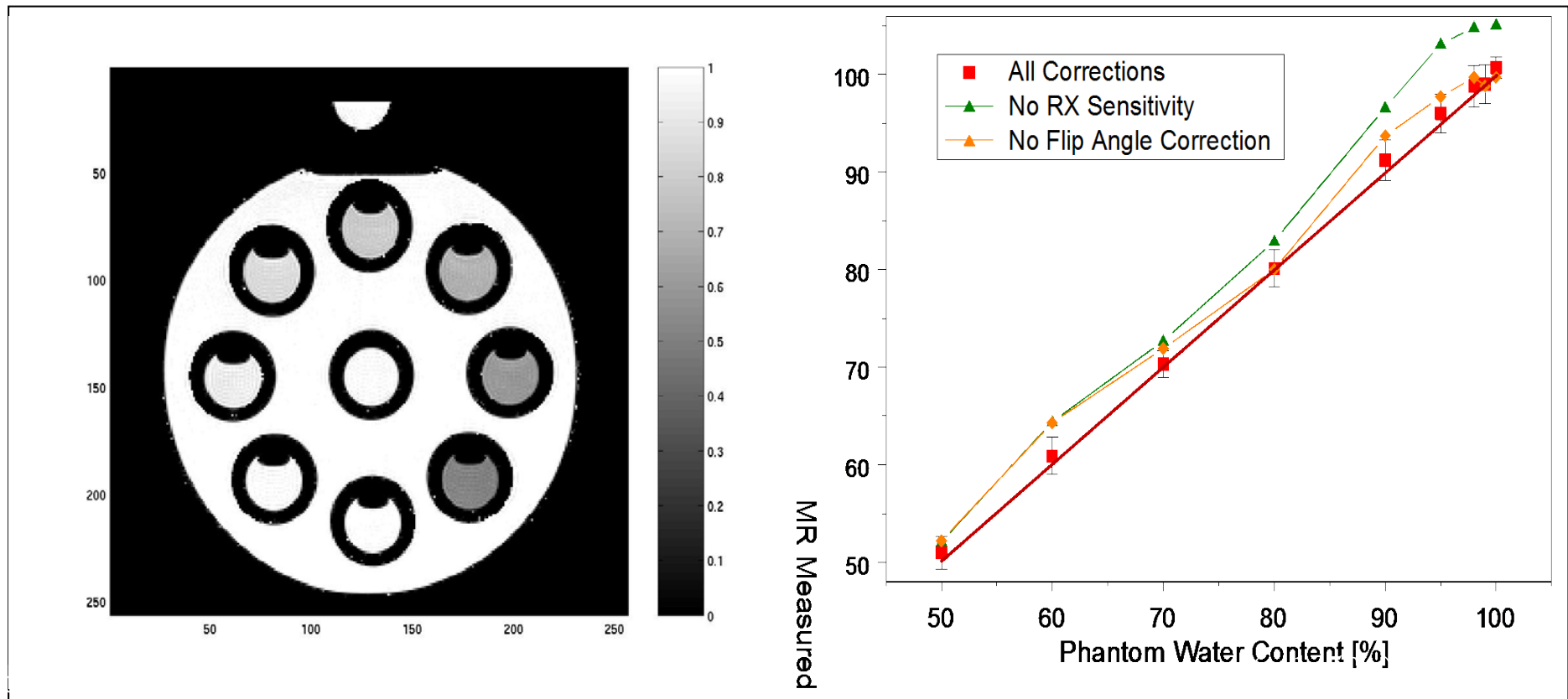


# Water Mapping

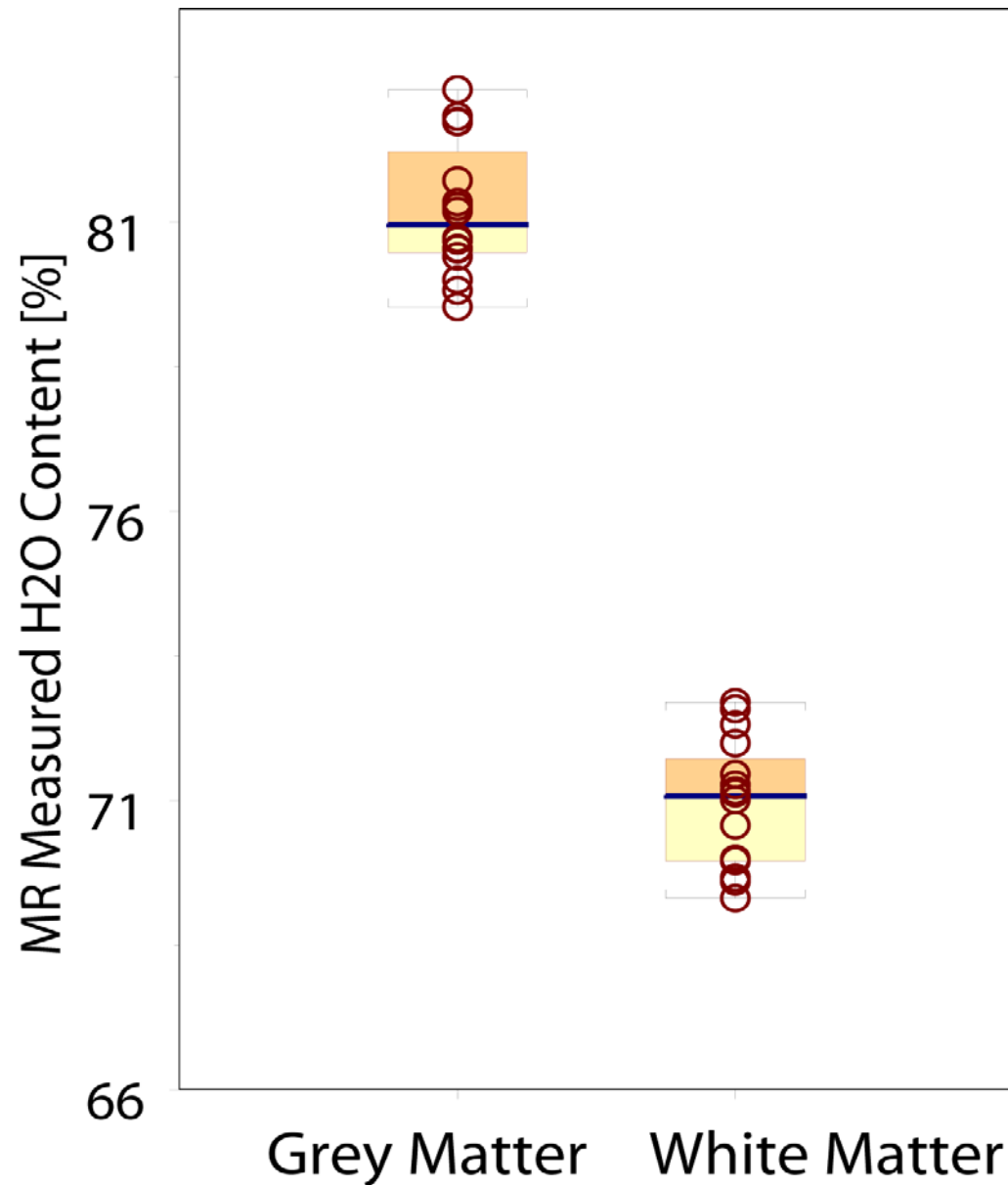


N. Jon Shah, Zaheer Abbas, Vincent Gras, Klaus Moellenhoff, Anca Oros-Peusquens

# Water Content Mapping

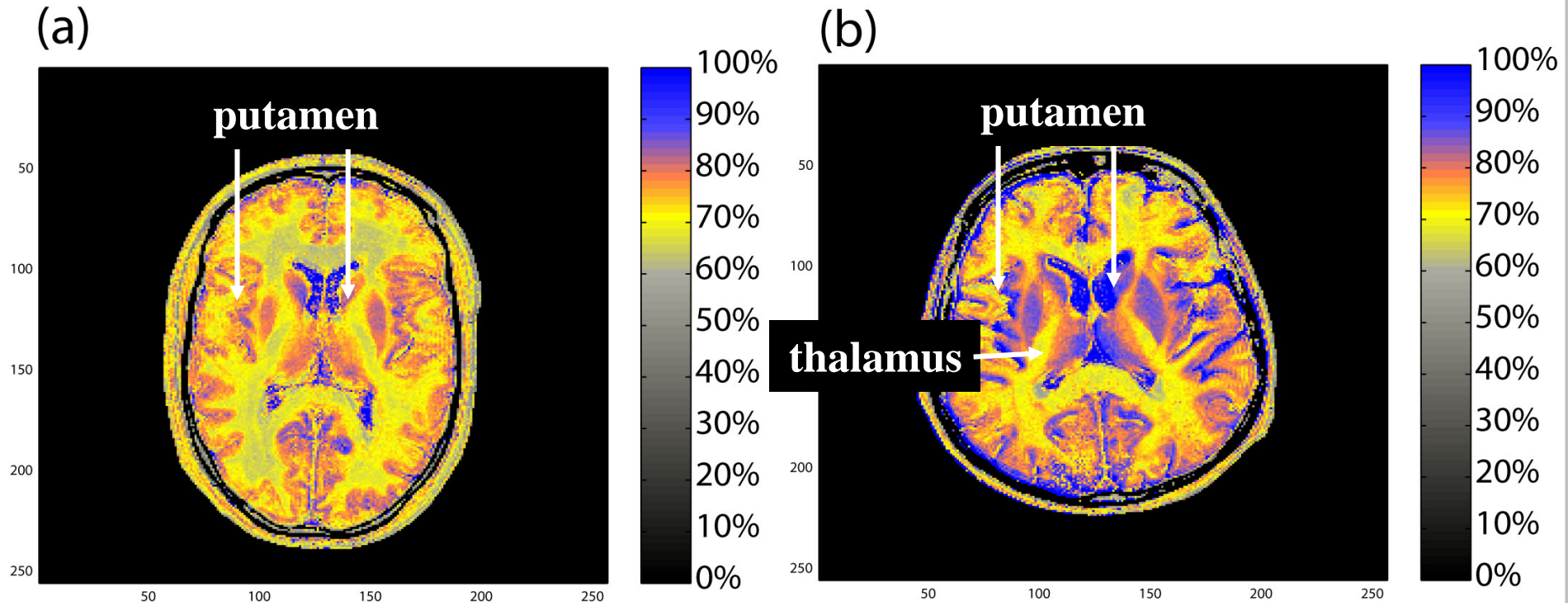


# Water Content in Grey/White Matter In Controls



Neeb et al., 2006a,  
NeuroImage 31 1156-  
1168

# Water Content Mapping @ 1.5T

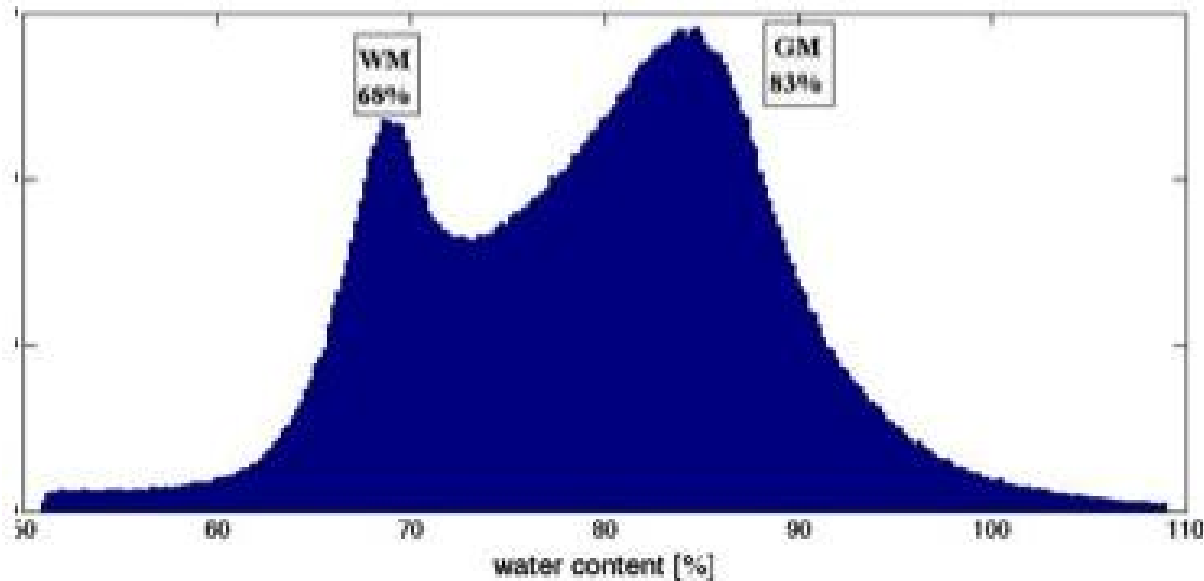


**Hepatic Encephalopathy grade HE 0      Hepatic Encephalopathy grade HE II**

Shah et al., US Patent No.: 6,803,762  
 Shah et al., NeuroImage: 2001 14(5): 1175-85  
 Steinhoff et al., Magn. Reson. Med.: 46(1) 131-140 2001  
 Zaitsev et al; Magn. Reson. Med.: 49(1) 1121-1132 2003  
 Shah et al., German Patent No.: 10028171

Shah et al., Hepatology: 2003 38: 1219-26  
 Neeb H, Shah NJ. Magn Reson Med. 2006 56(1):224-9.  
 Neeb H, Zilles K, Shah NJ. NeuroImage. 2006 31(3):1156-68.  
 Neeb H, Zilles K, Shah NJ. NeuroImage. 2006 29(3):910-22.  
 Shah et al., NeuroImage: NeuroImage 2008 41(3):706-17

# Test-retest stability

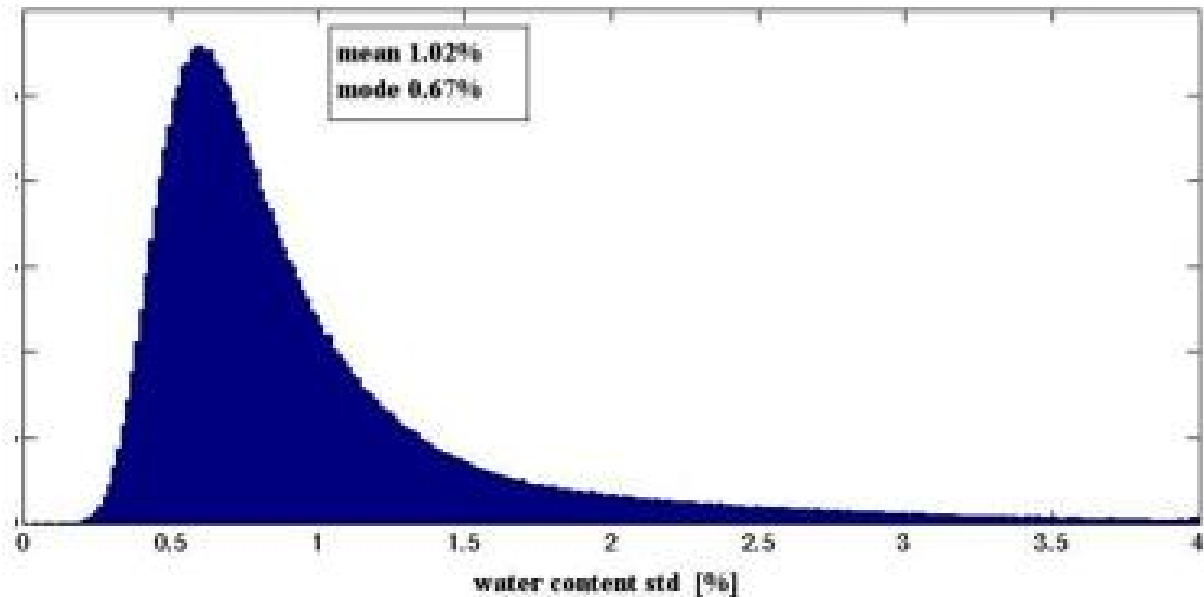


12 measurements

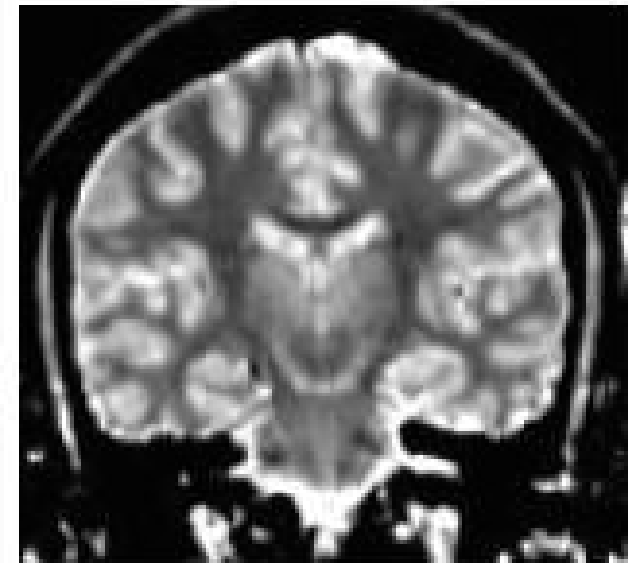
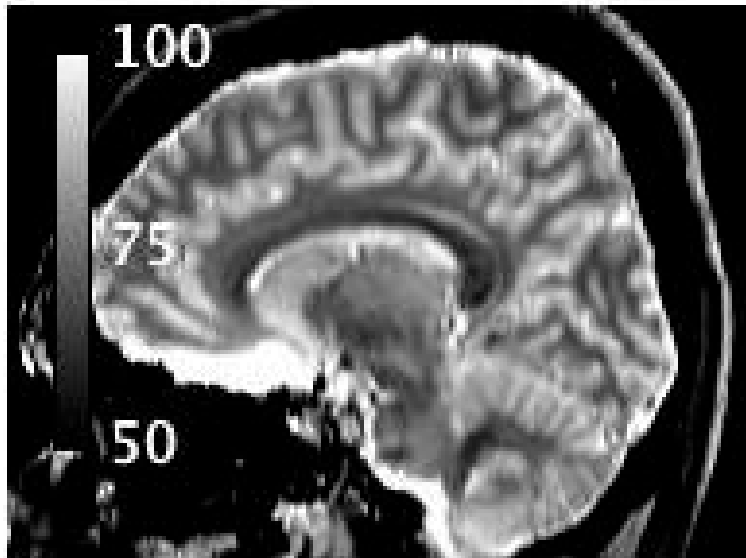
Repositioning

SD of mean  
values: **0.3%**

Voxel-based SD:  
**1%** mean value

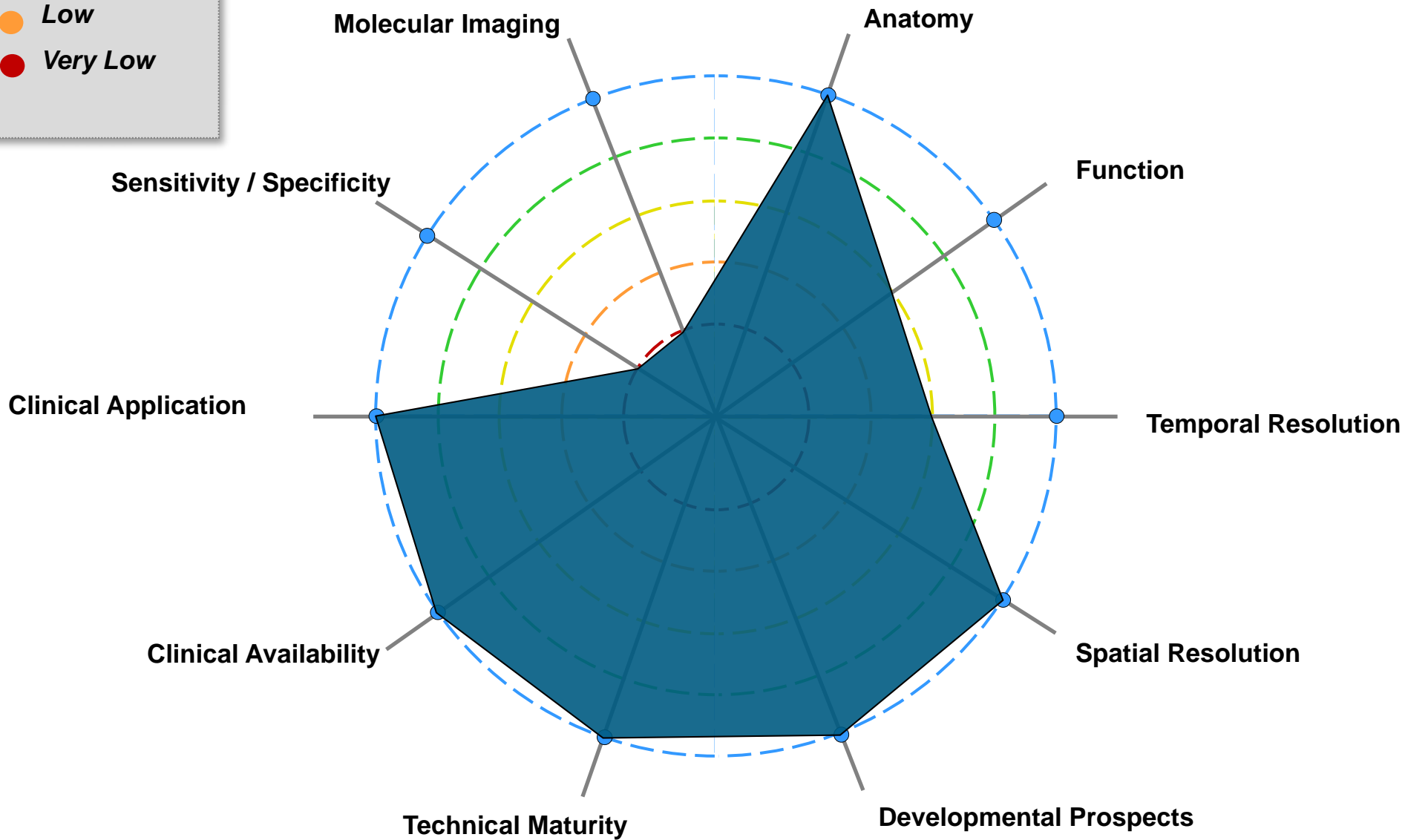


# Test-retest stability: mean water content map



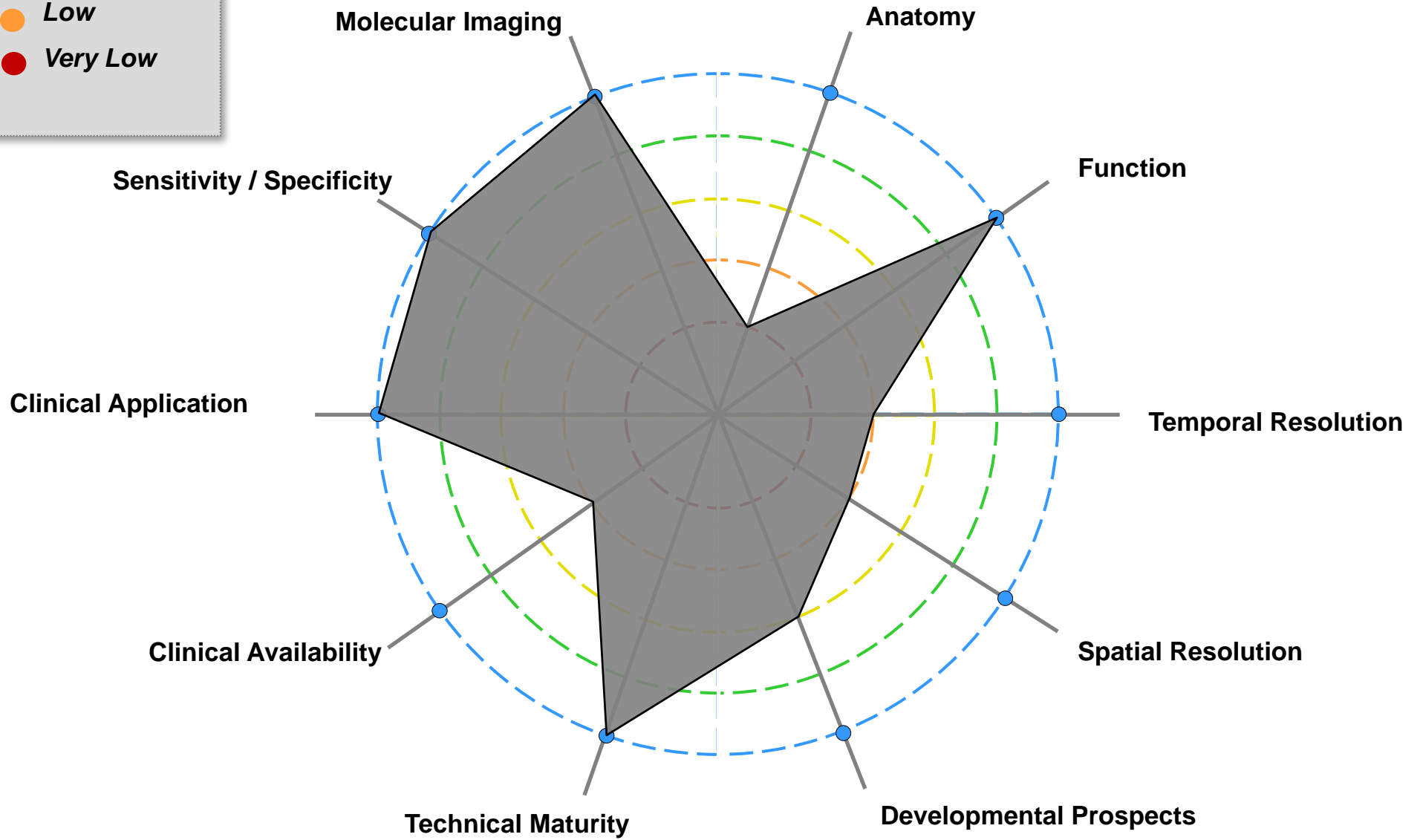
Exquisite anatomical detail, e.g. brain stem, thalamus

# MRI

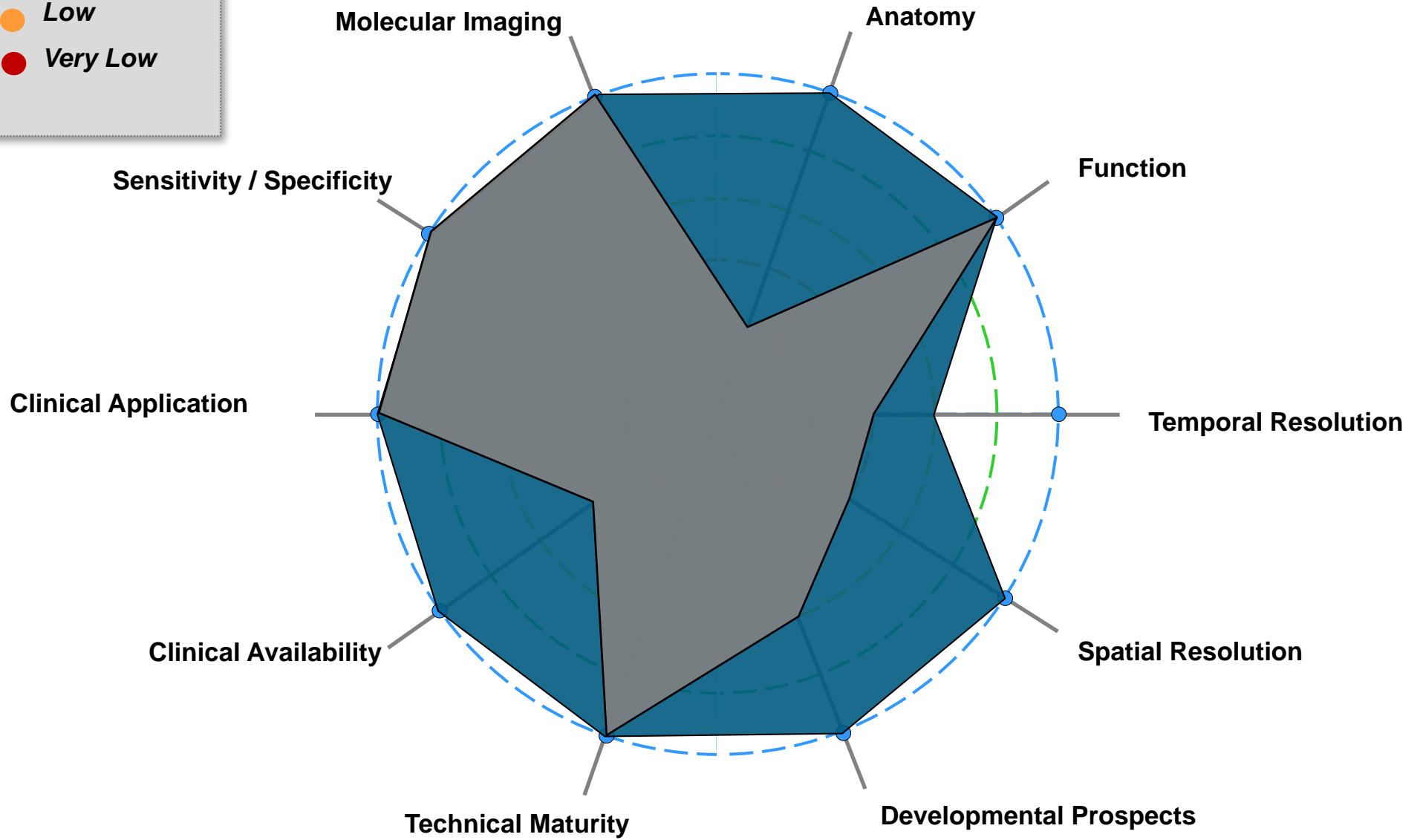




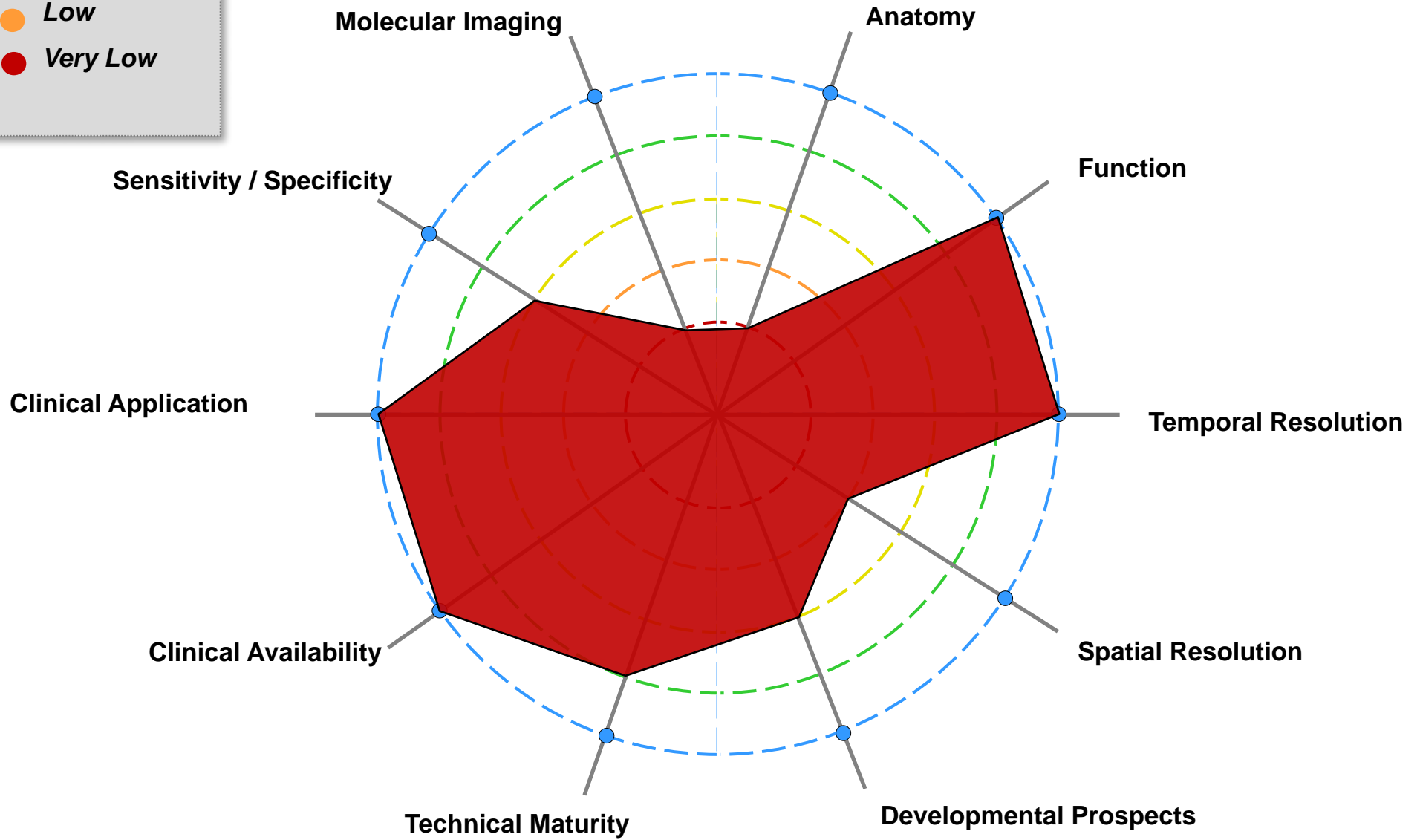
# PET



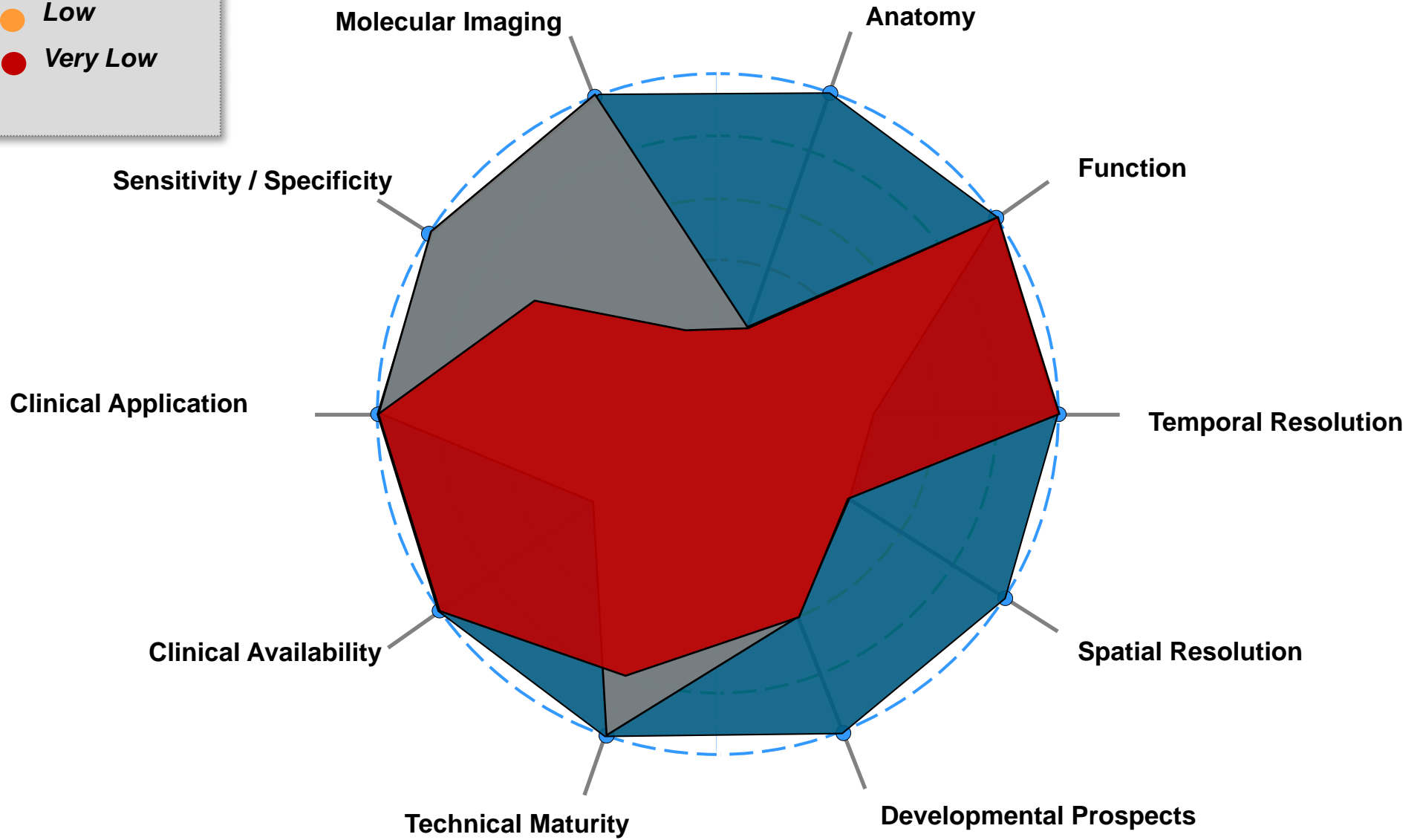
# MR-PET



# EEG

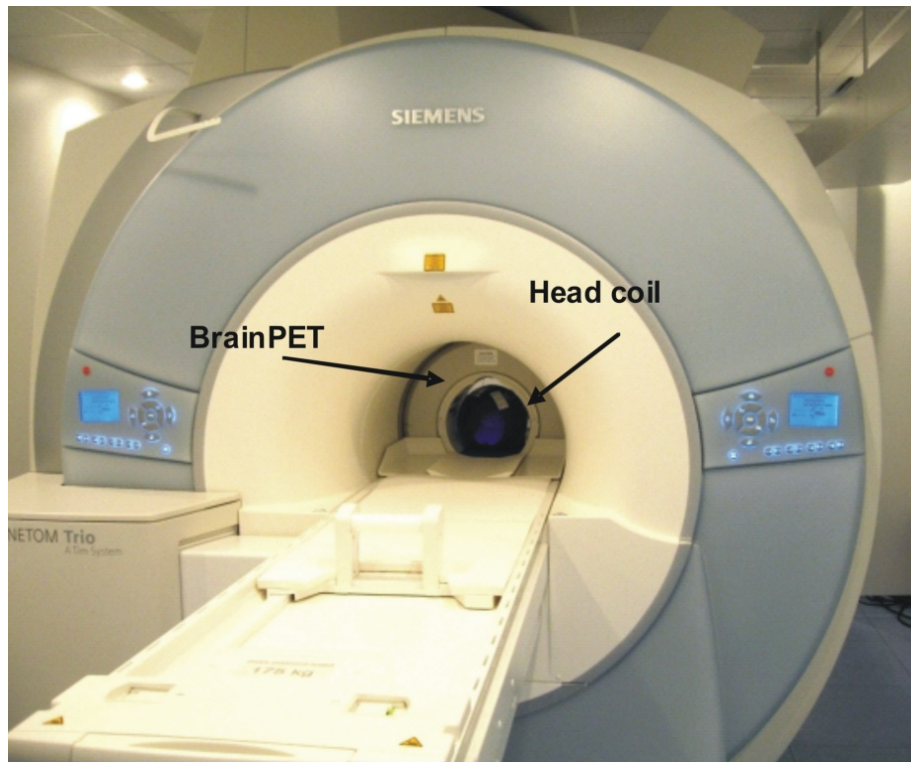


# MR-PET-EEG

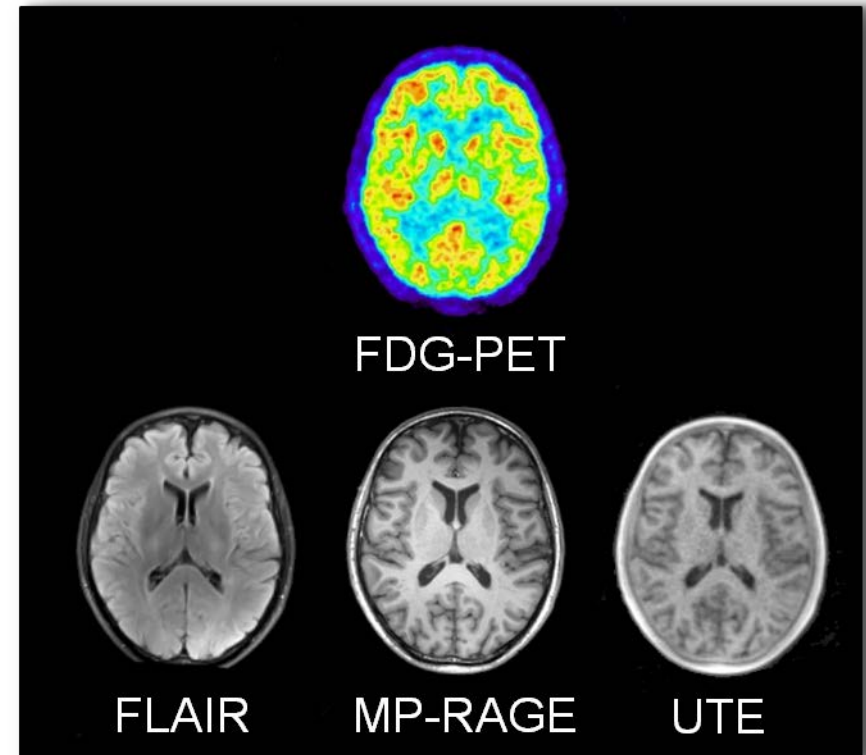


# 3T MR-PET

# Simultaneous 3T MR-PET Hybrid Measurements

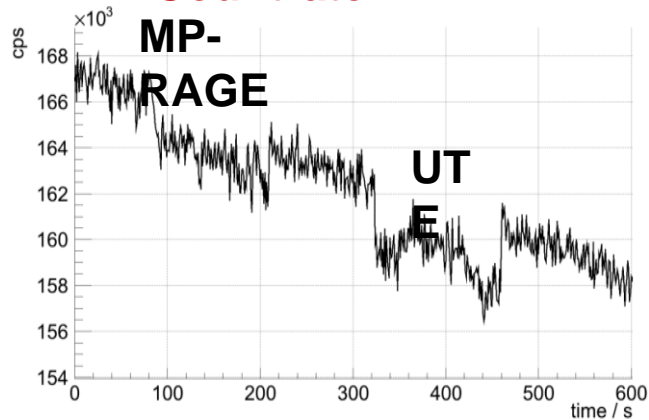


3T MR-PET hybrid scanner showing BrainPET and head coil

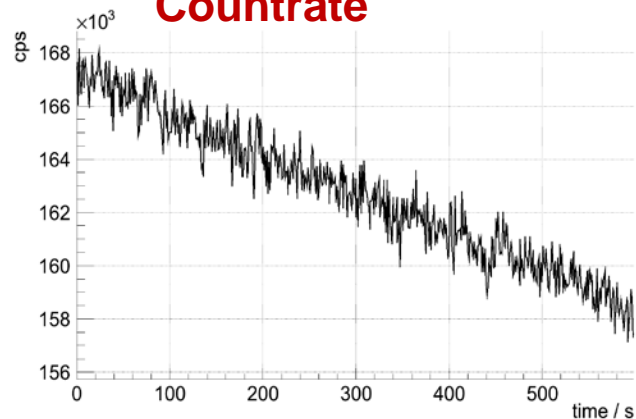


Simultaneous acquisition of  $^{18}\text{F}$ -FDG-PET and MR images

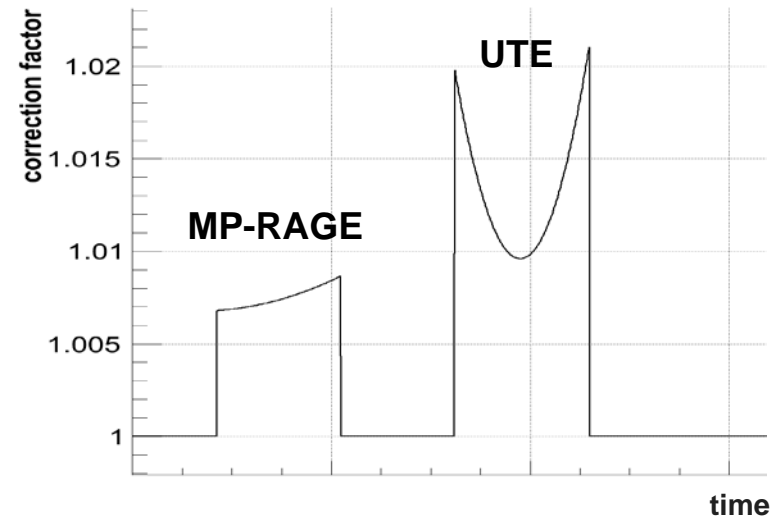
## Uncorrected Countrate



## Corrected Countrate



## Correction Factor



## Cross Calibration of the PET Scanners

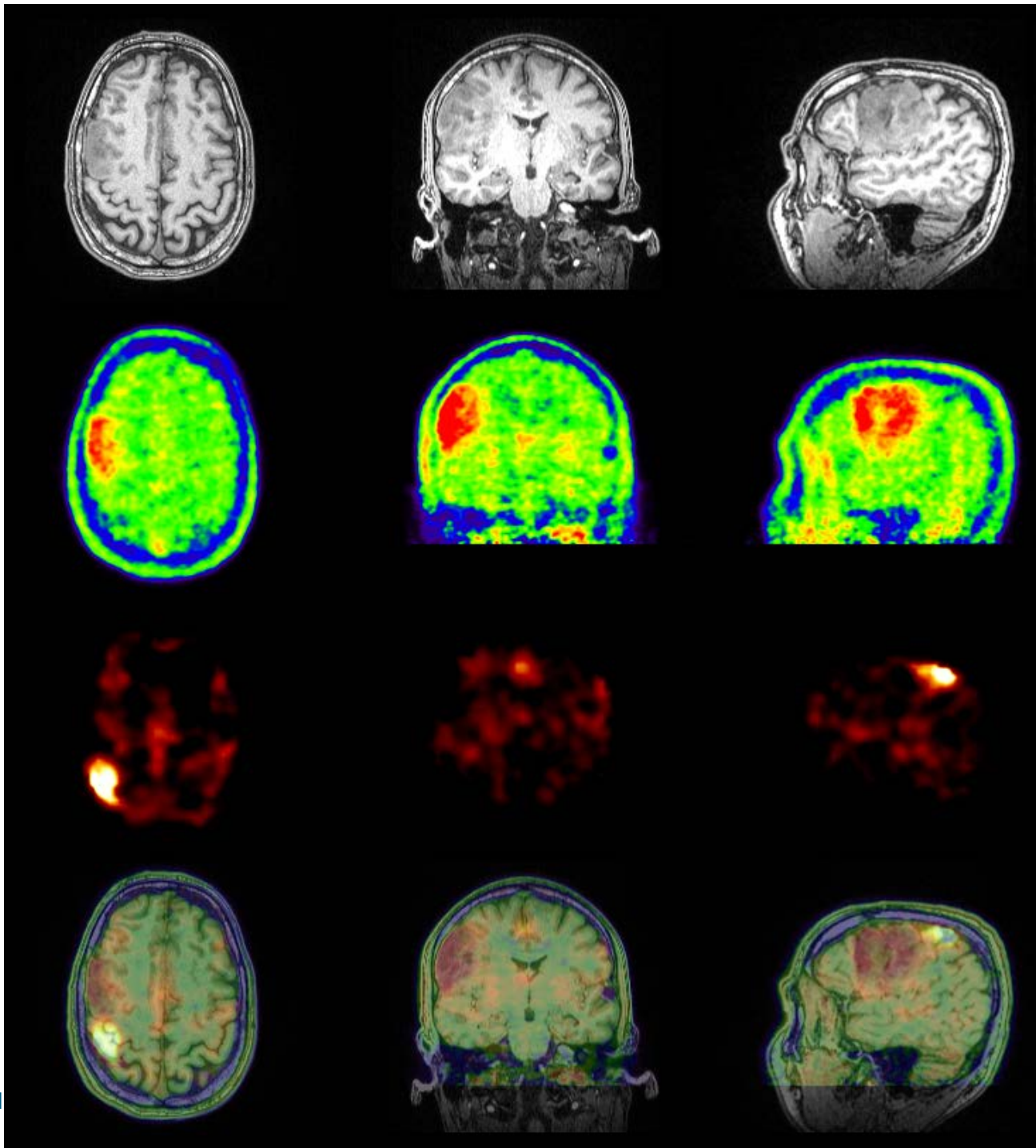
- Randoms Correction
- Scatter Correction
- Attenuation Correction of Head
- Normalisation of Crystal Efficiencies
- Deadtime and Pileup Correction
- Attenuation Correction of RF Coil
- MRI Interference Correction

Weirich, ..., Shah (2012) *IEEE Trans. Med. Imaging*

# Clinical Applications

K.-J. Langen et al.





T1 MPRAGE (6 min)

PET:  
[<sup>18</sup>F]-fluor-ethyl-tyrosine  
20 - 40 min p.i.

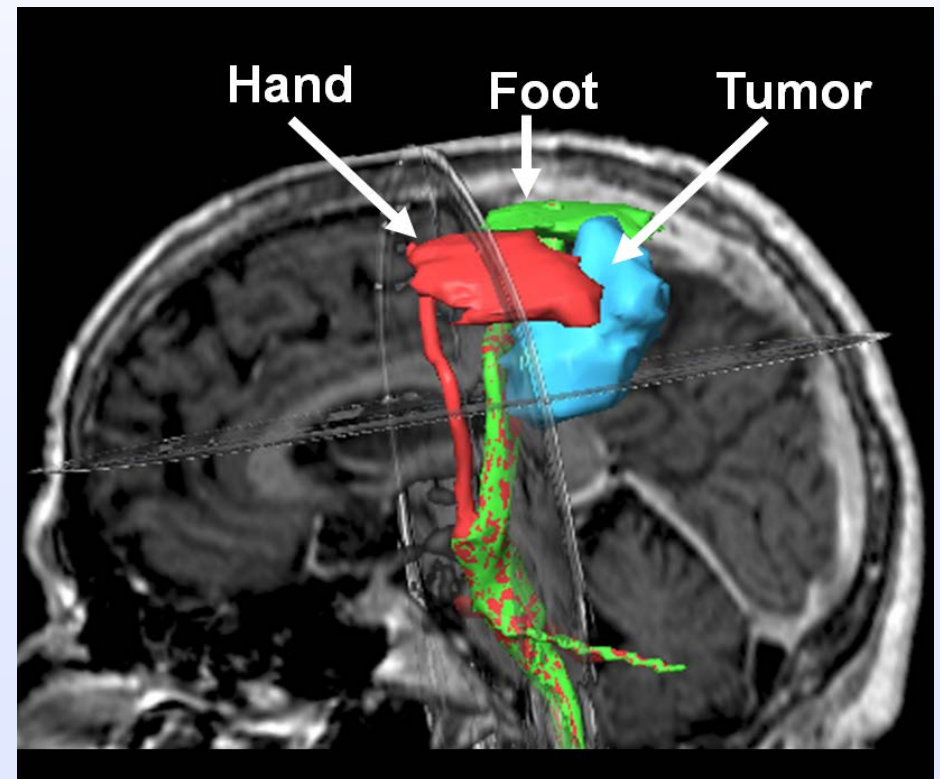
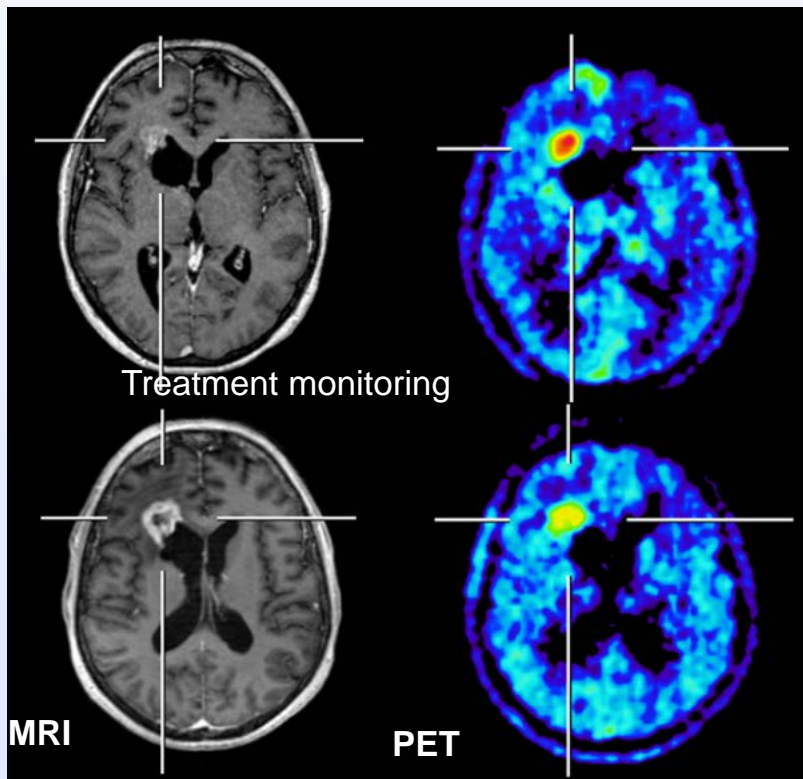
BOLD imaging:  
Finger tapping  
left hand

Fusion

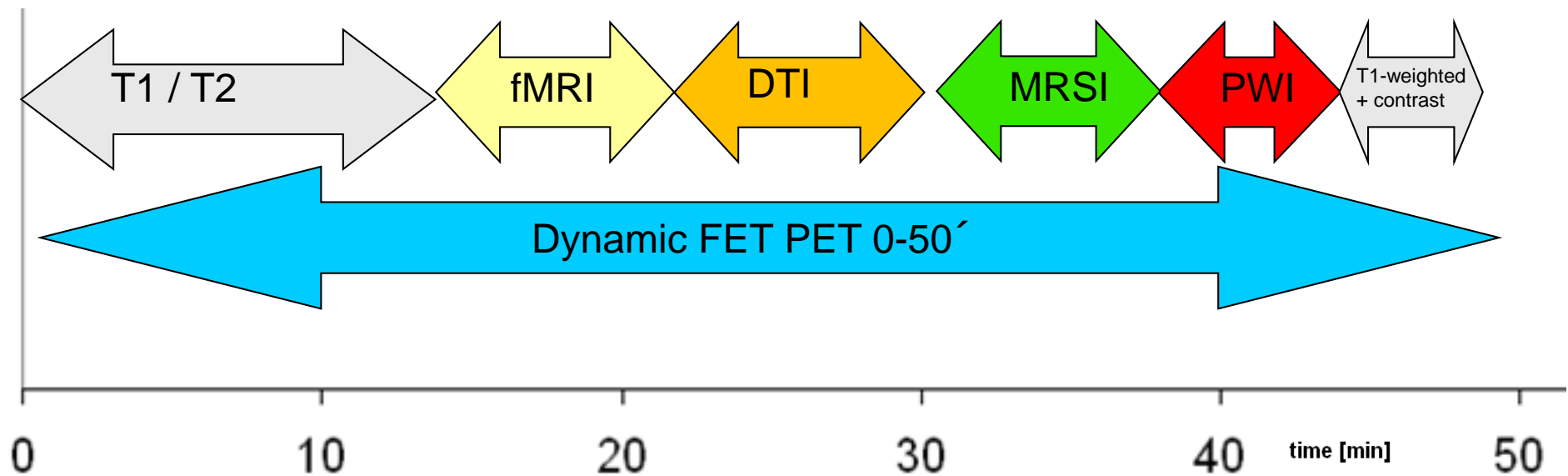
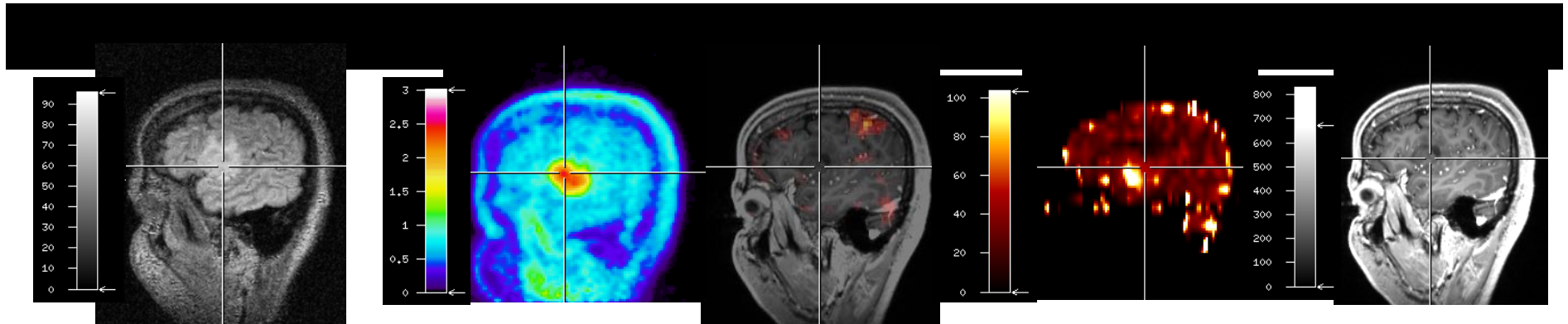
# Brain Tumours

$^{18}\text{F}$ - FET PET  
Clinical Studies

Hybrid  
MR-PET Imaging



# Hybrid MR-PET imaging



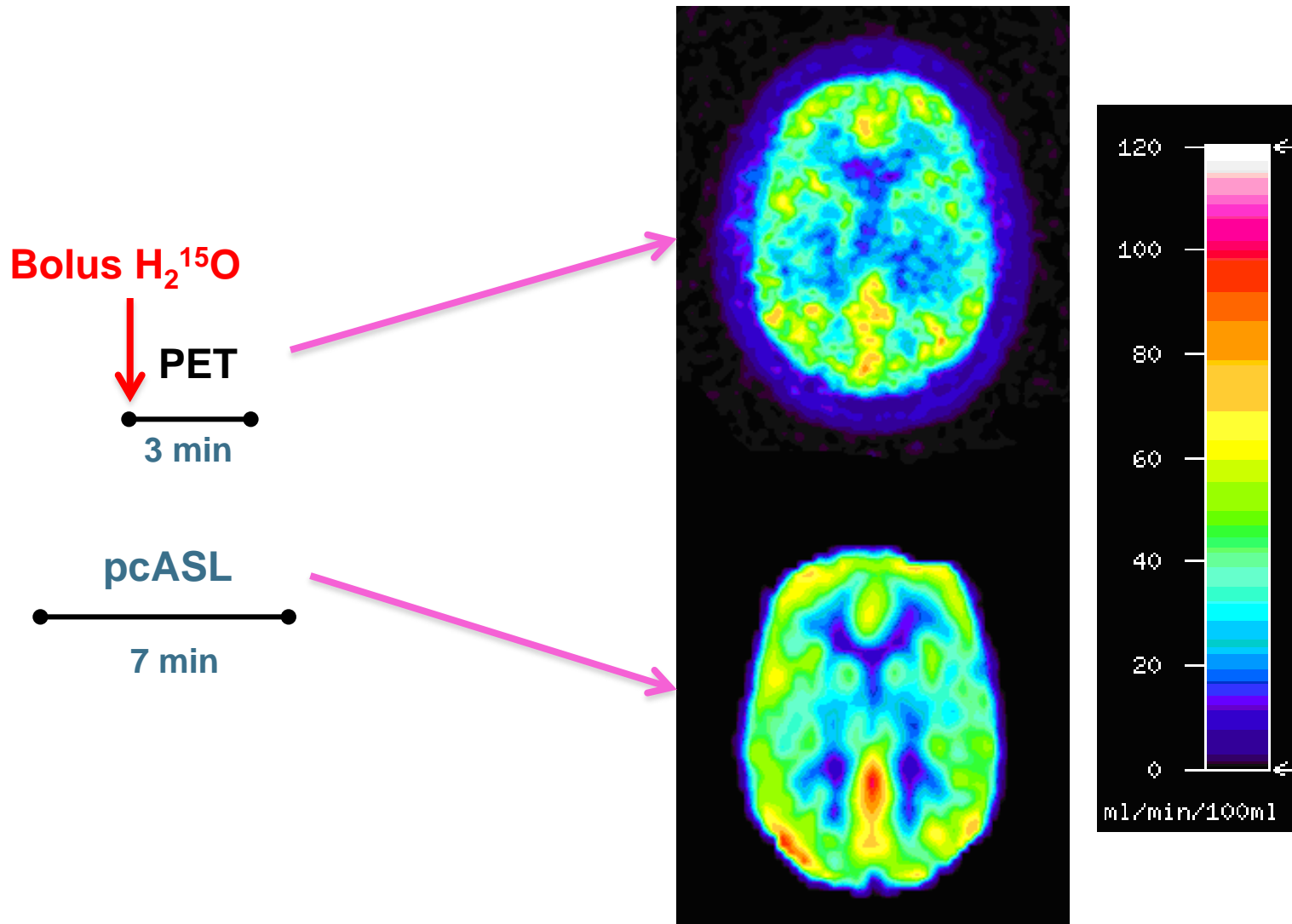
# CBF

## $^{15}\text{O}$ -Water PET

# Arterial Spin Labelling

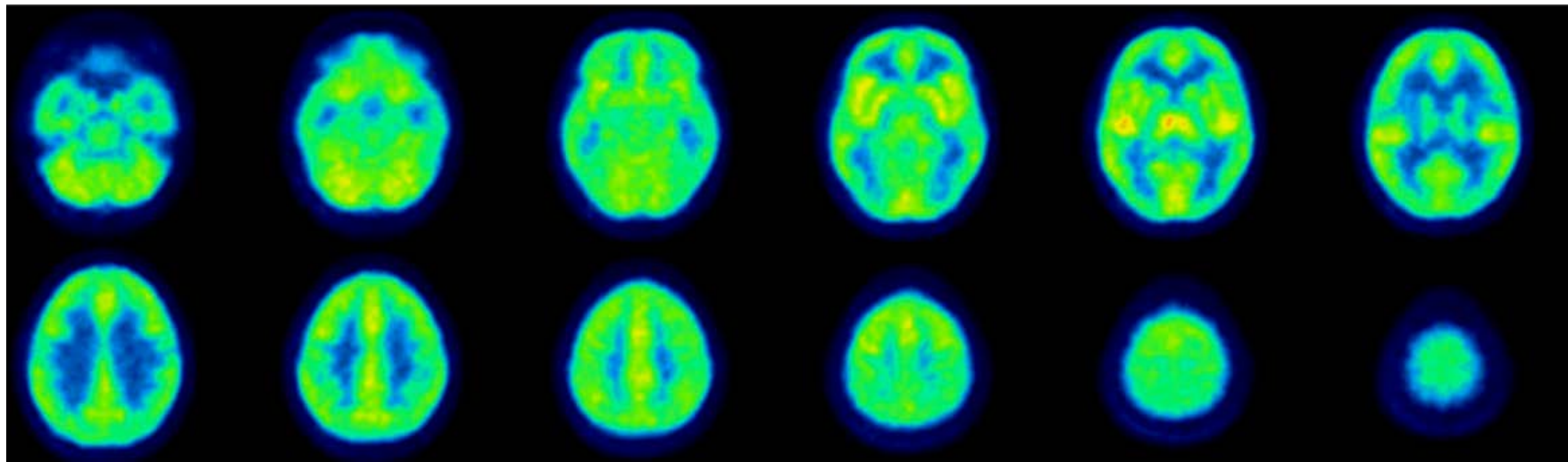
K. Zhang, H. Herzog et al.

# First Truly Simultaneous Comparison of CBF Assessed by $^{15}\text{O}$ -Water PET and ASL

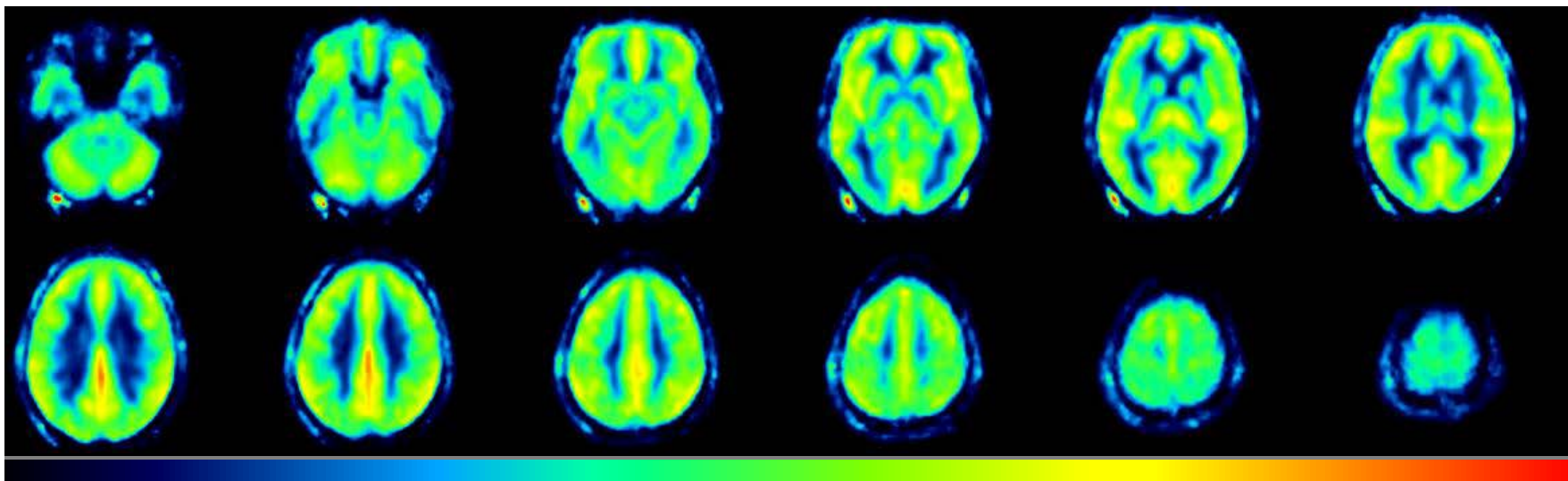


Sequence courtesy of Tom Okell and Peter Jezzard (FMRIB, Oxford)

# Averaged Results



**PET-CBF**



**ASL-CBF**

0.0

ml/min/100ml

120.0

**Averaged CBF images (n=10) after normalisation to MNI space**

**ASL:  $51.9 \pm 7.1$  ml/100g/min**

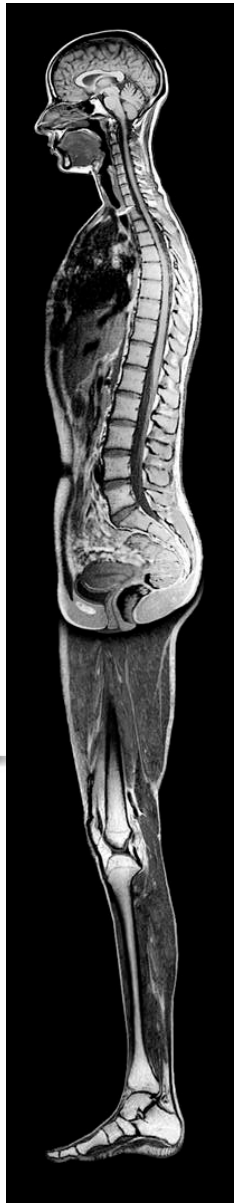
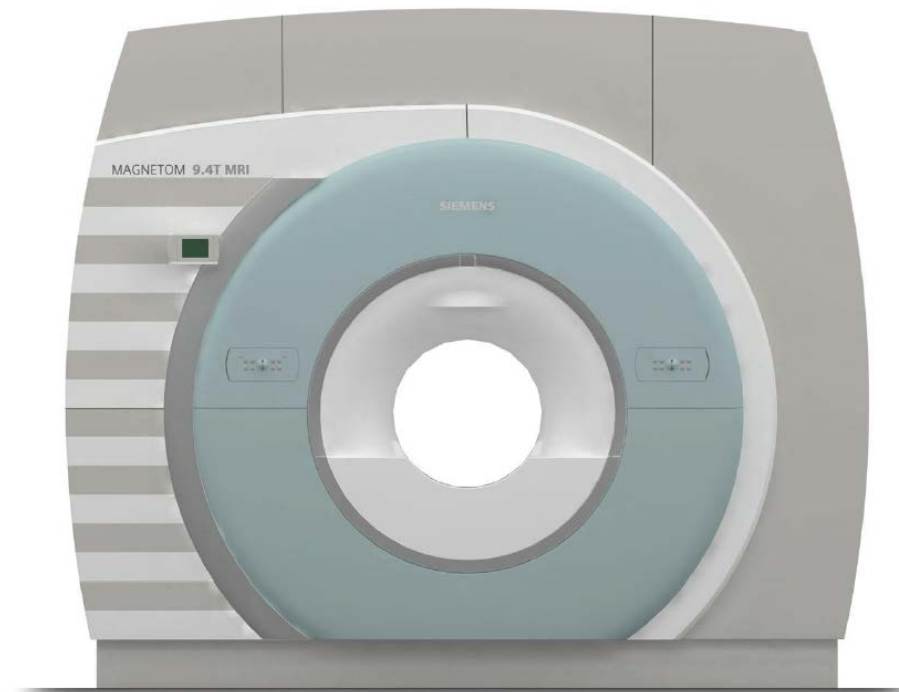
**PET:  $48.1 \pm 9.9$  ml/100g/min**

# High-Field MRI



## 9.4T Whole-Body Scanner in Jülich

- 60 cm patient bore
- TQ-engine gradient coil
- 50 cm FoV
- Magnet weight: 57 tonnes
- 870 tonnes of iron shielding
- 3.70 m length
- Stored energy: 182.0 MJ
- Length of wire: 750 km



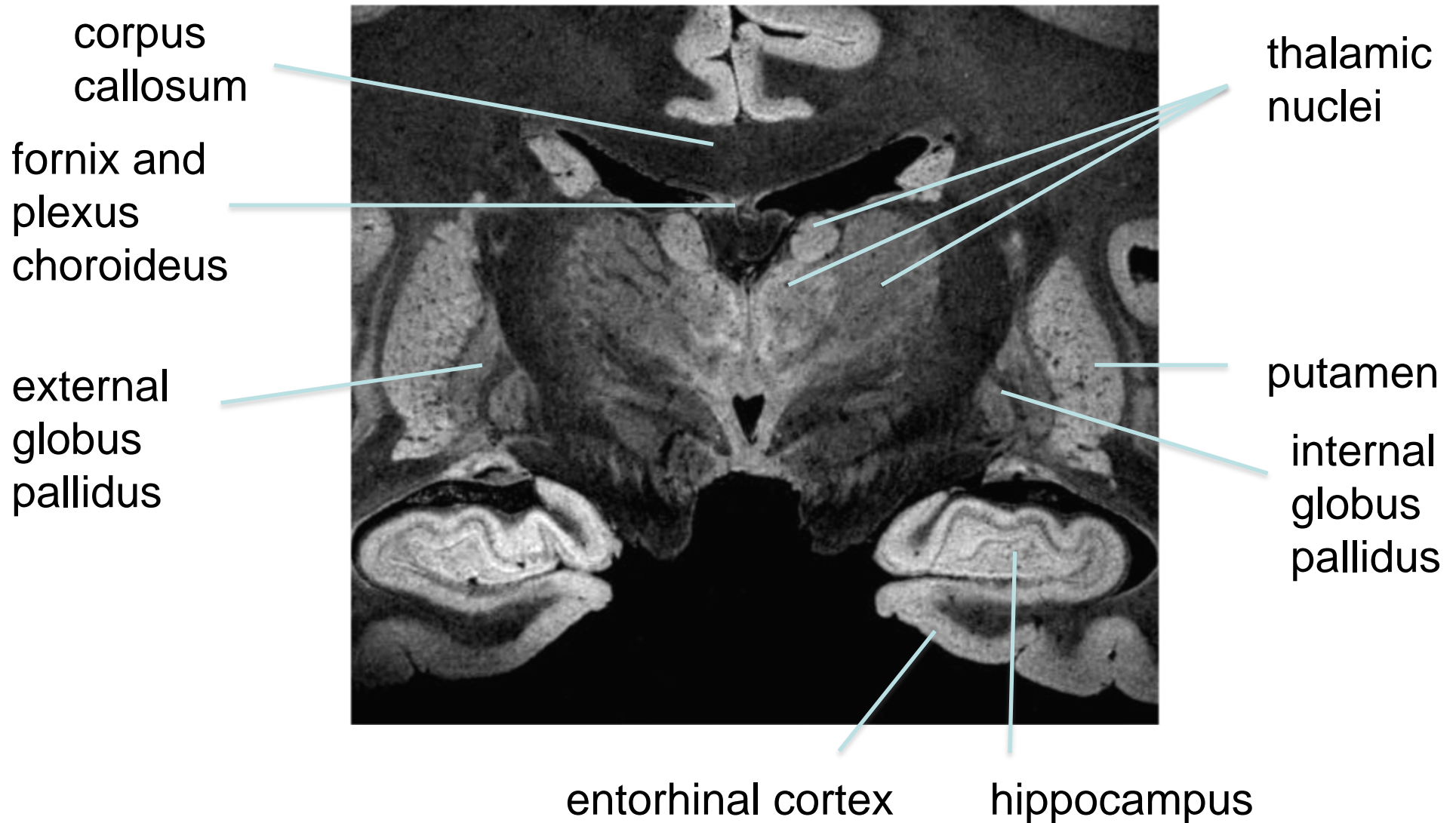
**Complete with Hybrid PET Capability!**



# Structural Imaging

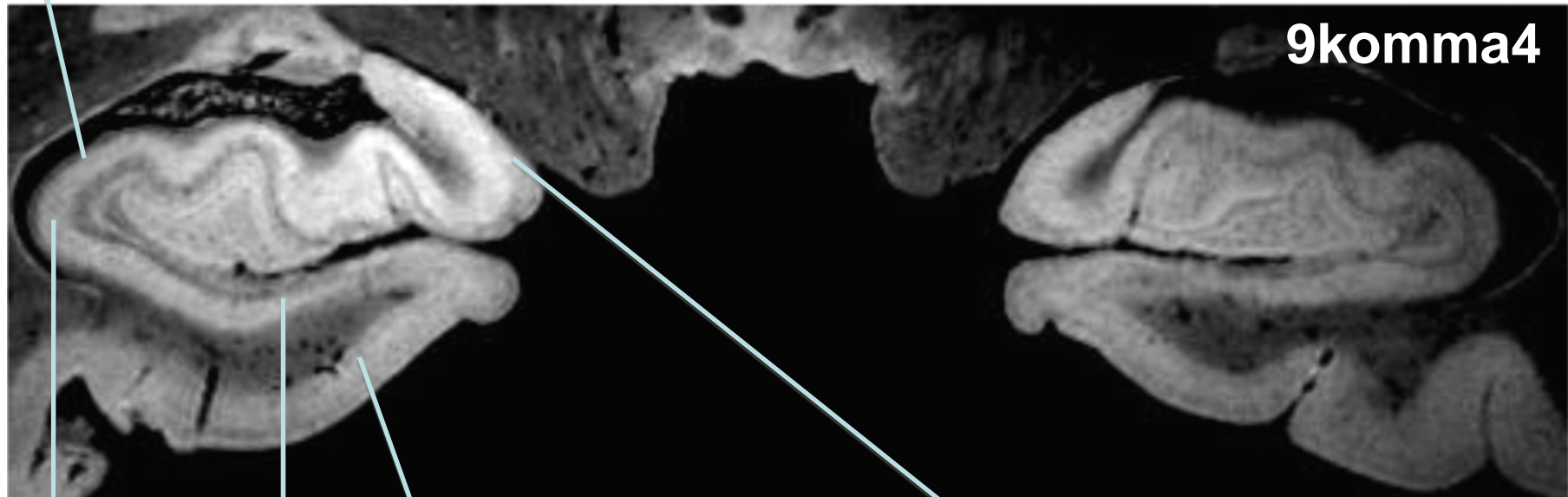
A.-M. Oros-Peusquens, J. Lindemeyer et al.

# Hippocampus, thalamus



# Anterior Hippocampus

cornu ammonis (CA2)



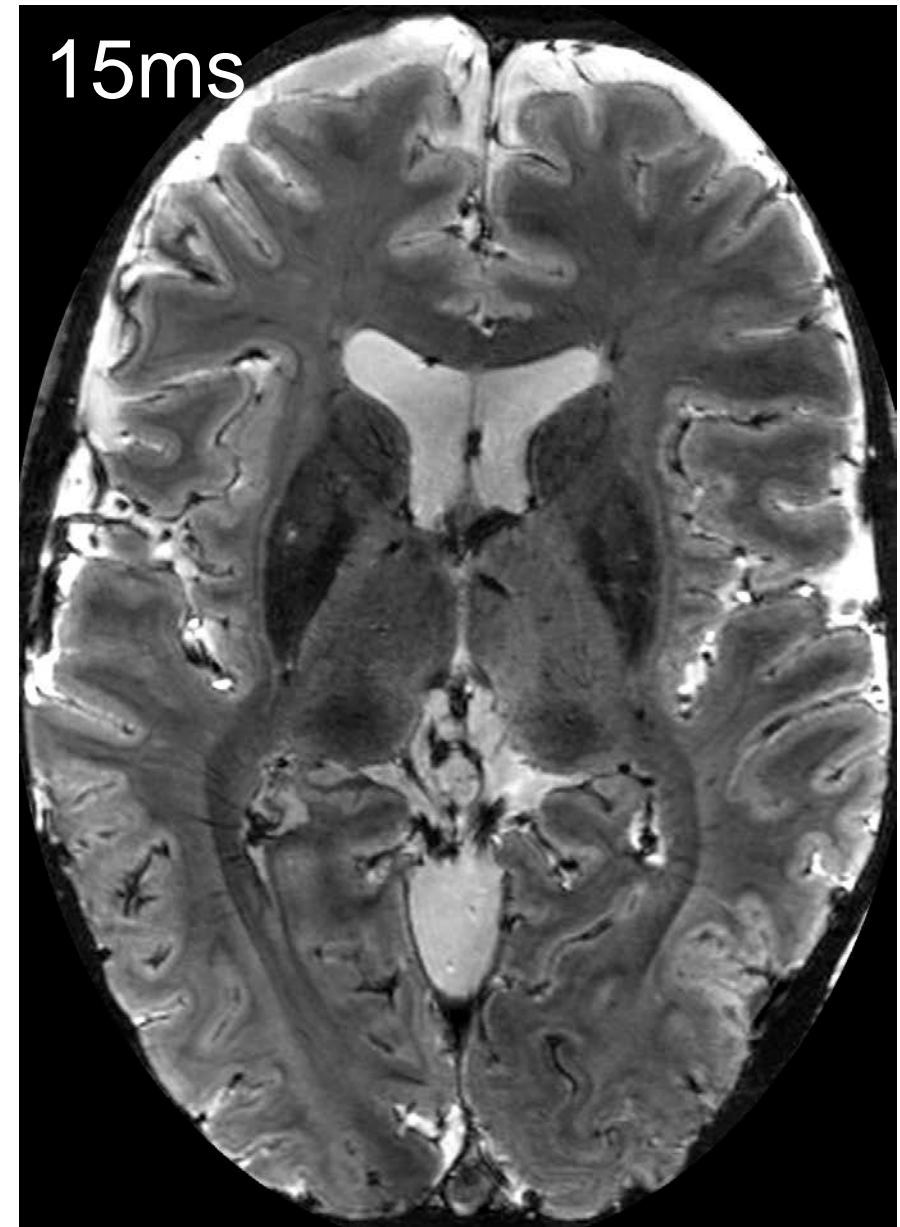
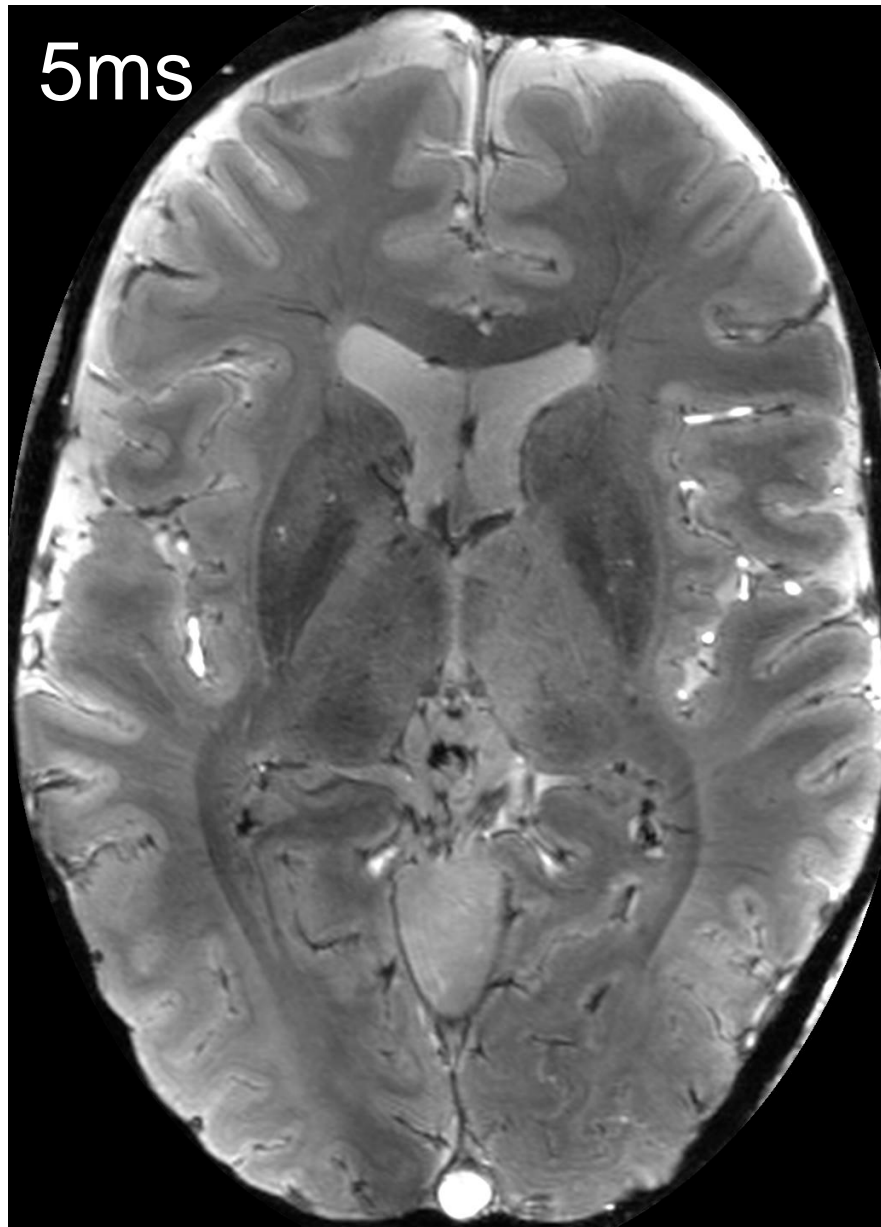
cornu ammonis  
(CA1)

subiculum

entorhinal cortex

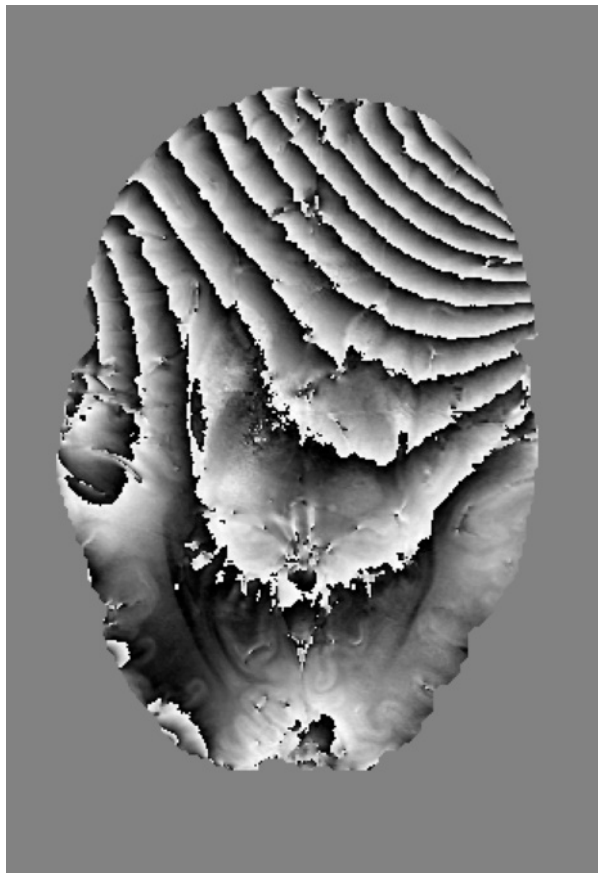
gyrus dentatus

# Structural imaging at 9.4T with $(0.5\text{mm})^3$ resolution

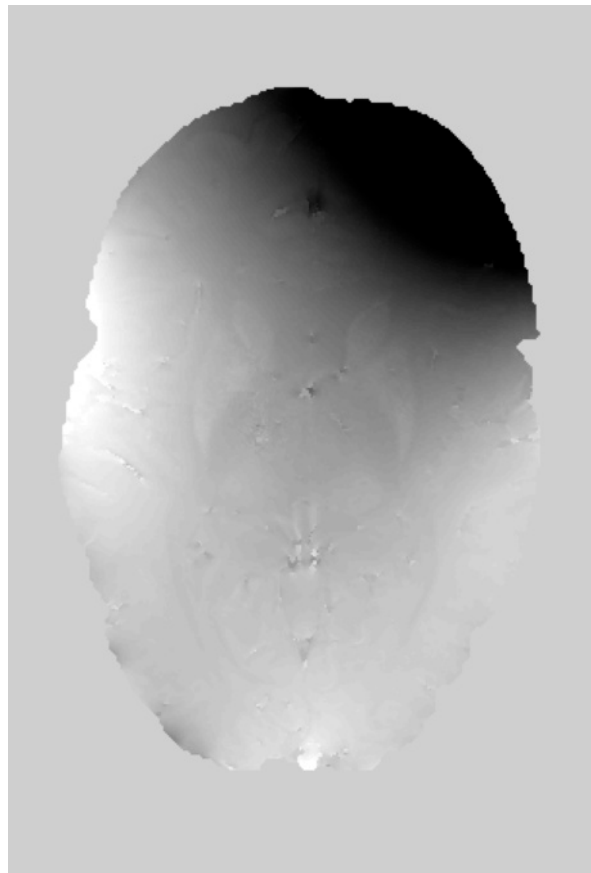


# Phase Imaging at 9.4T (*in vivo*)

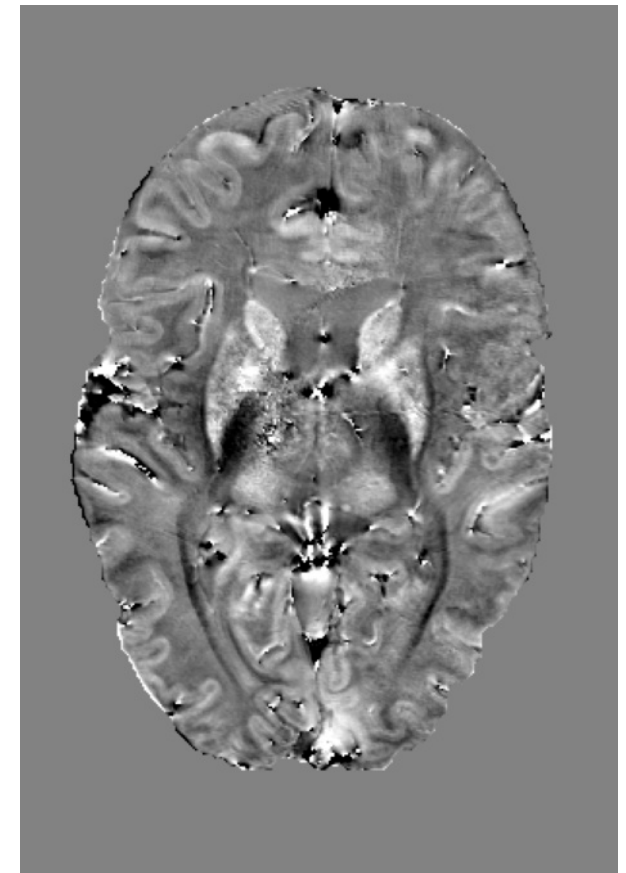
Phase



Unwrapped Phase (URSULA)  
→ Fieldmap



Background-Corrected  
(MUBAFIRE)

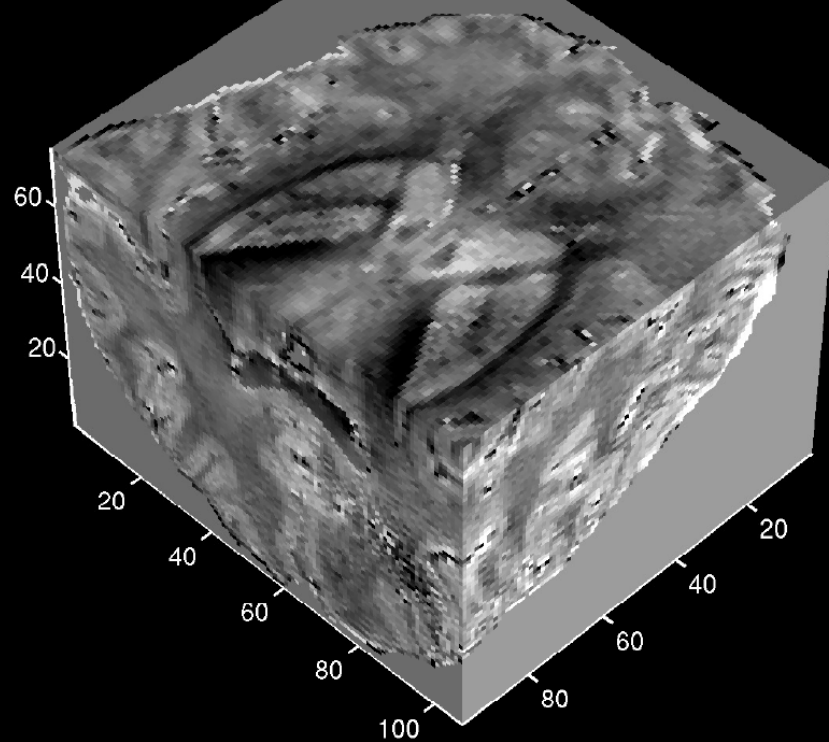


*GRE 0.5mm isotropic, slab-selective excitation (central brain)*

## Phase Imaging at 9.4T (*in vivo*)

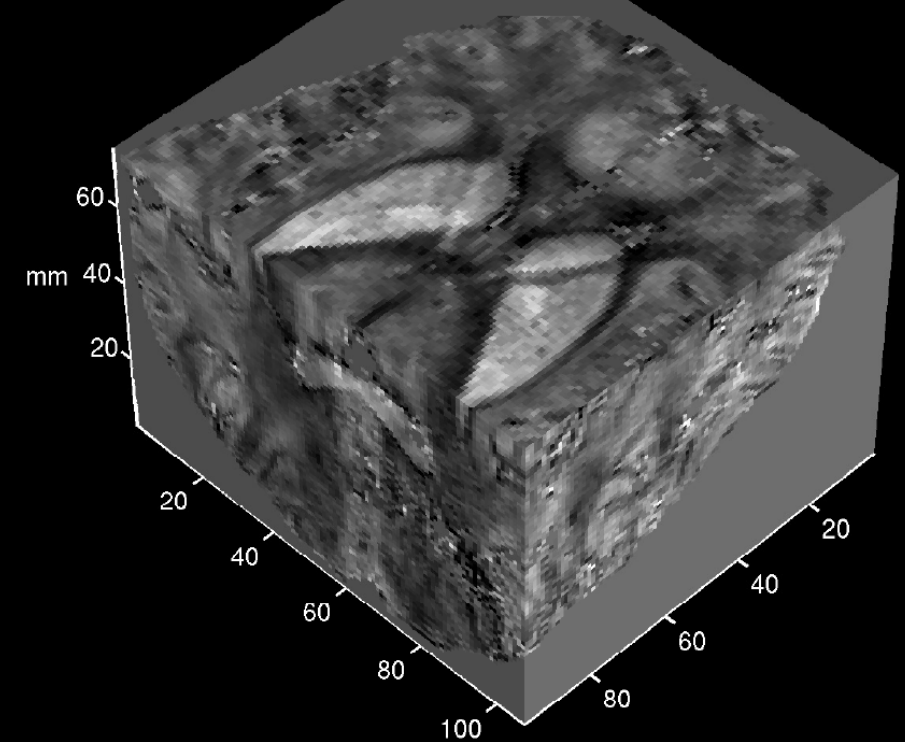
- Background-corrected field and susceptibility at 1mm isotropic, whole brain coverage

Field



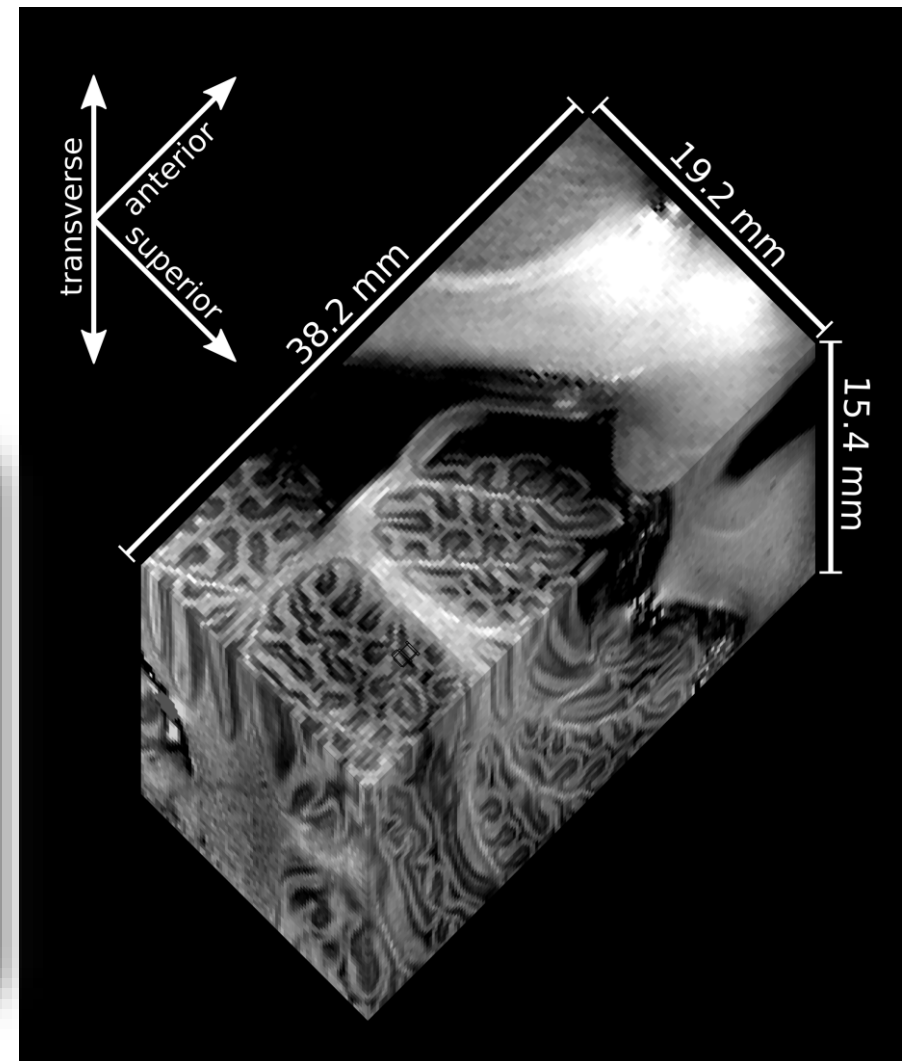
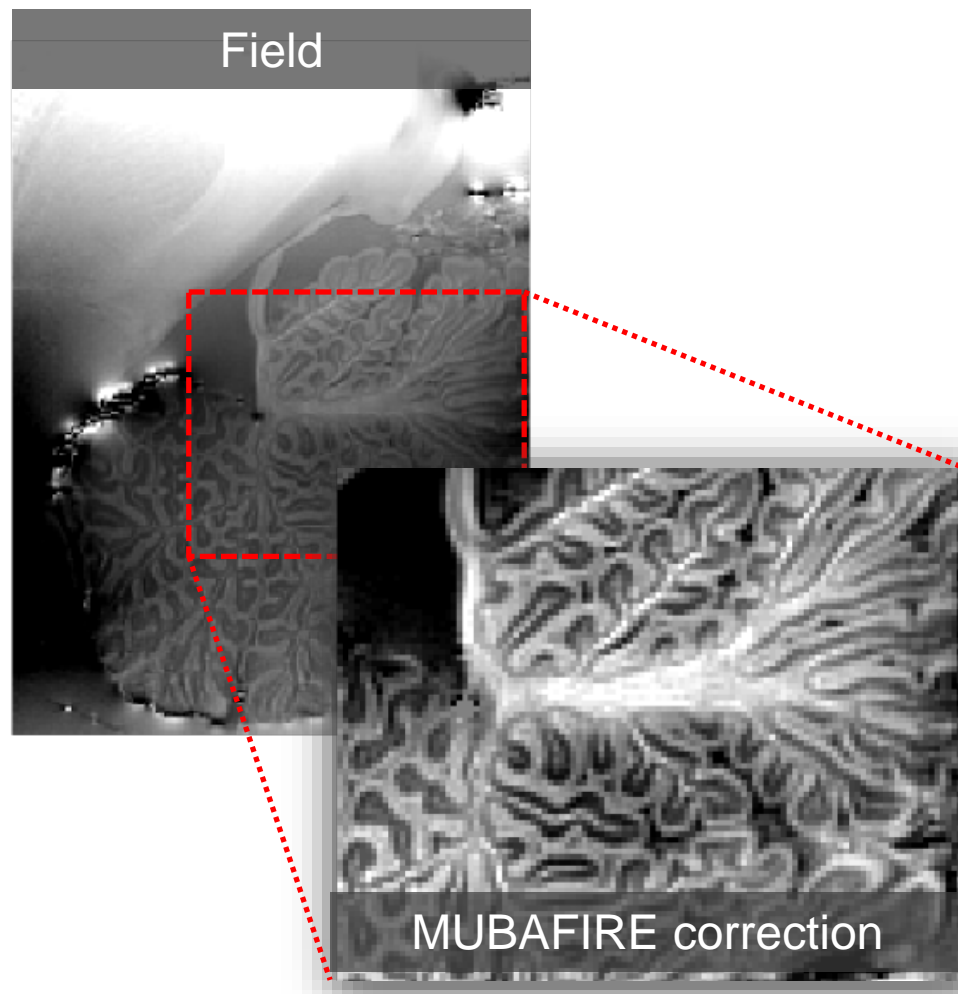
Feldverschiebung [Hz]

Susceptibility



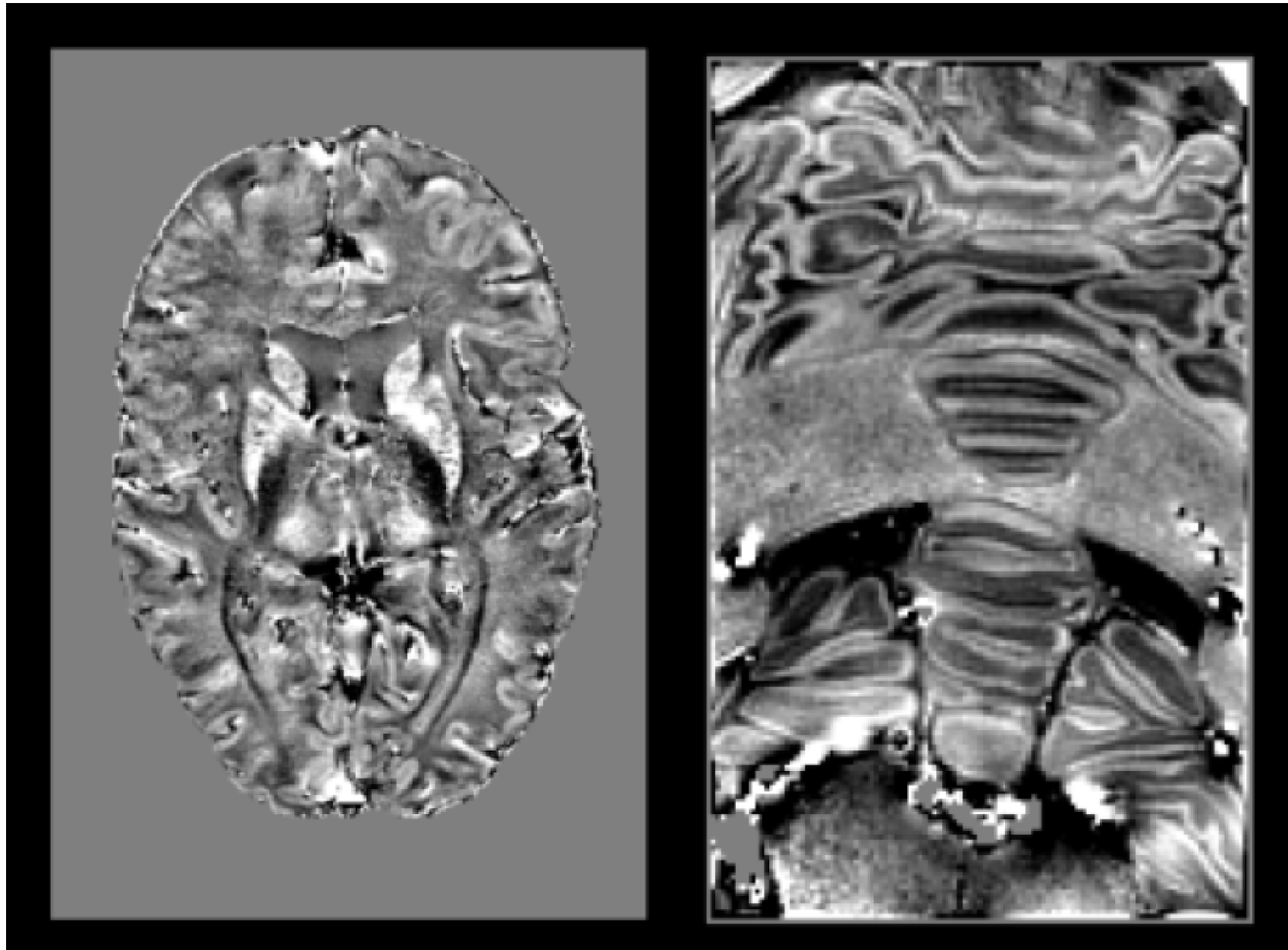
Suszeptibilität ( $\chi$ ) [ppm]

## Phase Imaging at 9.4T (*post mortem*)



- Phase of cerebellum at 0.24mm isotropic

# Towards MR histology...

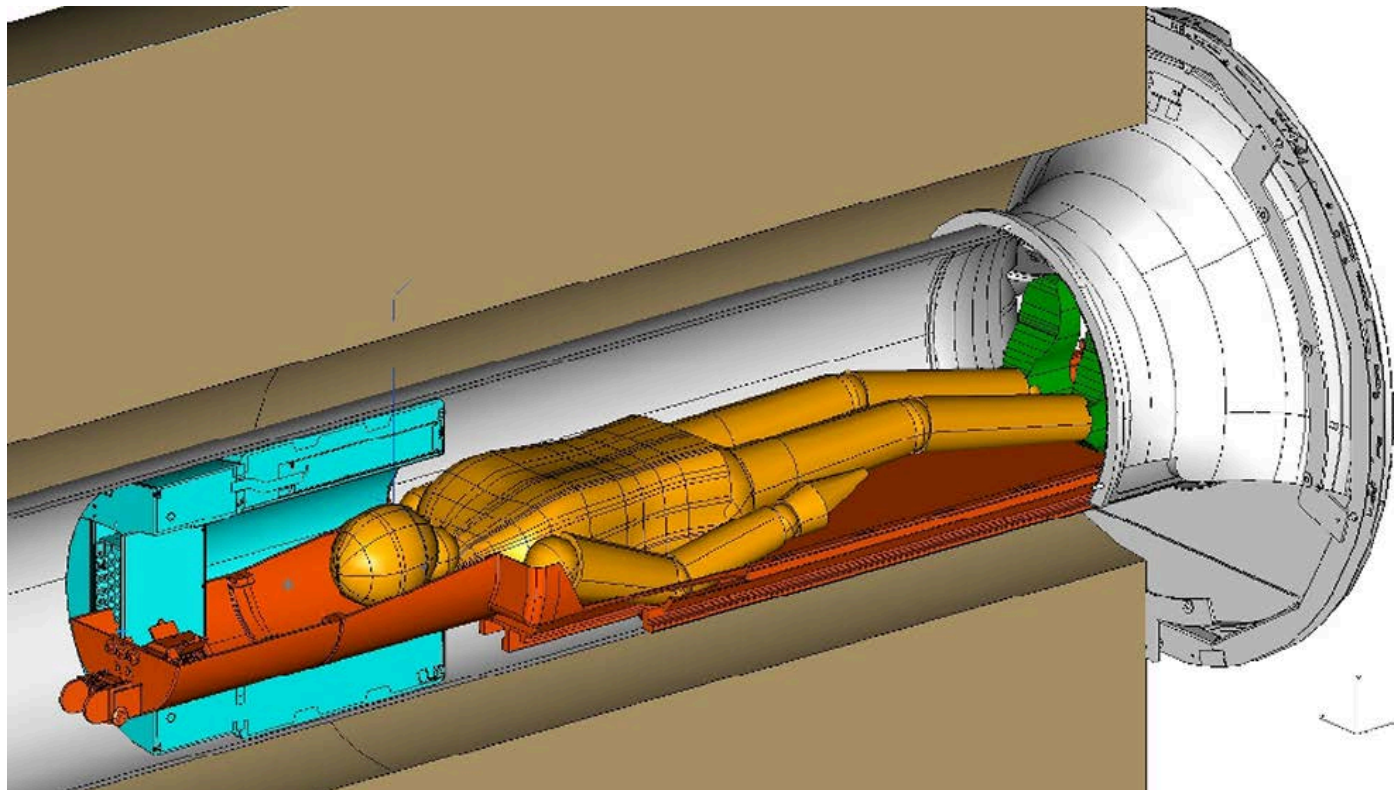




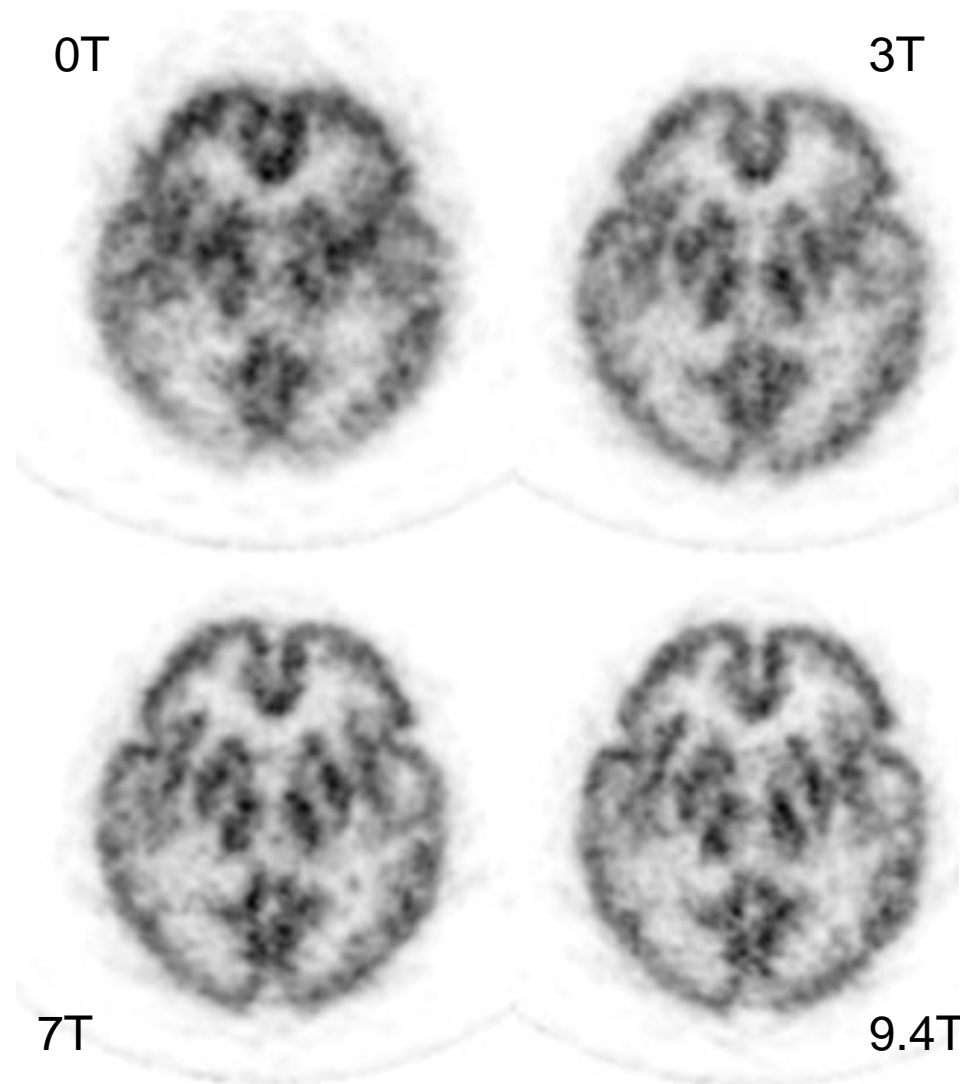
# MR-PET @ 9.4T

N. J. Shah, H. Herzog, C. Weirich et al.

# 9.4T MR-PET



# Positron Range at 9.4T using the “Iida” Brain Phantom

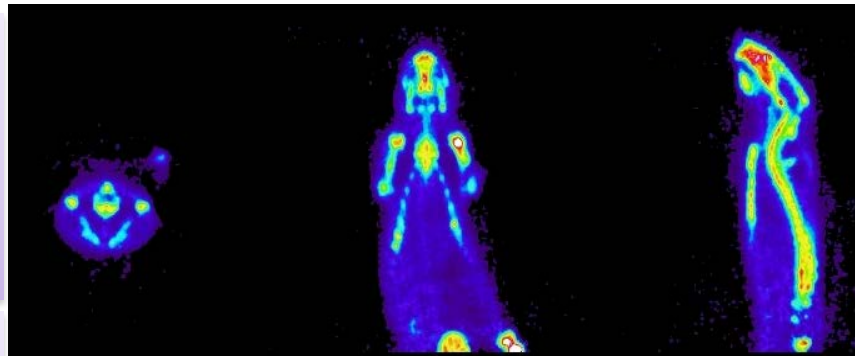


Shah et al. (2014)  
*PLoS ONE*  
(in press)

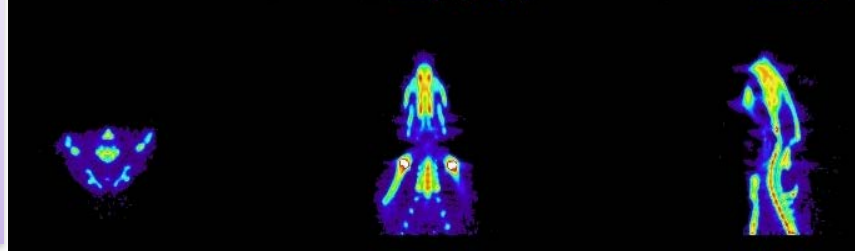
Polymer brain phantom filled with  $^{120}\text{I}$

# Rat Bone Scan with $^{18}\text{F}$ -Fluoride

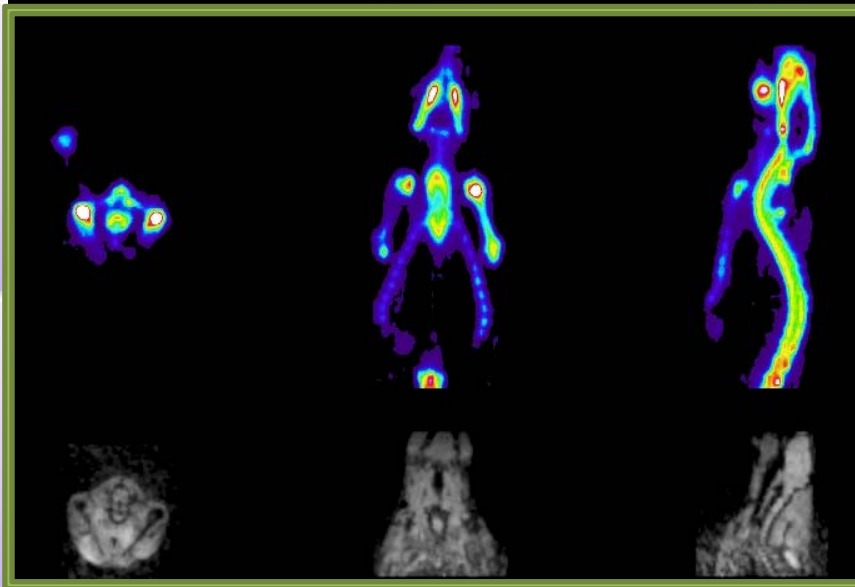
BrainPET @ 3 T



Inveon PET



BrainPET @ 9.4 T



T2-weighted MRI @ 9.4  
T

Shah (2014) *BS&F* (in press)

# Opportunities for Hybrid MR-PET

## ... MRI

- ⇒ Higher spatial resolution (structural imaging)
- ⇒ Higher functional (BOLD) contrast => columnar resolution fMRI?
- ⇒ Better image quality (contrast)
- ⇒ Non-proton MRI and spectroscopy

## ... PET

- ⇒ Partial volume correction with MRI
- ⇒ Attenuation correction with MRI
- ⇒ Motion correction with MRI (navigator echoes)

## ... Hybrid MR-PET

- ⇒ Patient / volunteer compliance: 2 scans in 1 (at 3T and 9.4T)
- ⇒ **Metabolic imaging (e.g. FDG +  $^{17}O$  +  $^{31}P$  +  $^{23}Na$  + MP-RAGE)**
- ⇒ Accurate receptor density mapping
- ⇒ Novel paradigms for brain function

# Opportunities – Metabolic Imaging

## ... Sodium

⇒ Na / K Pump

⇒ Disturbances of the pump often leads to cell death

⇒ Intra vs extracellular sodium with TQF

## ... Phosphorus

⇒ Energy metabolism of the cell

⇒ In vivo pH

## ... Oxygen

⇒ Intimately involved in metabolism!

⇒ .....

## ... Glucose

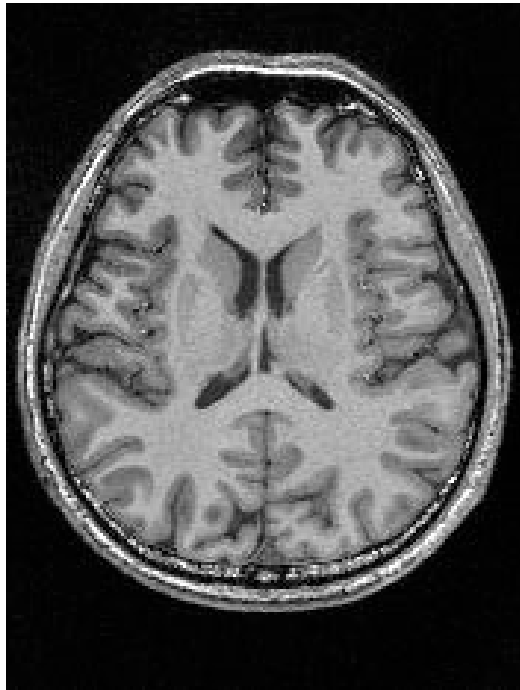
⇒ Energy substrate of the brain

⇒ FDG PET

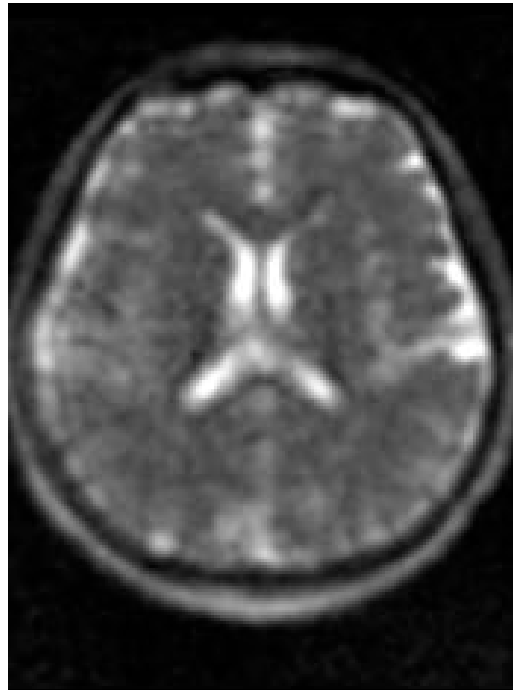
# Sodium Imaging

S. Romanzetti, D. Fiege, N. J. Shah et al.

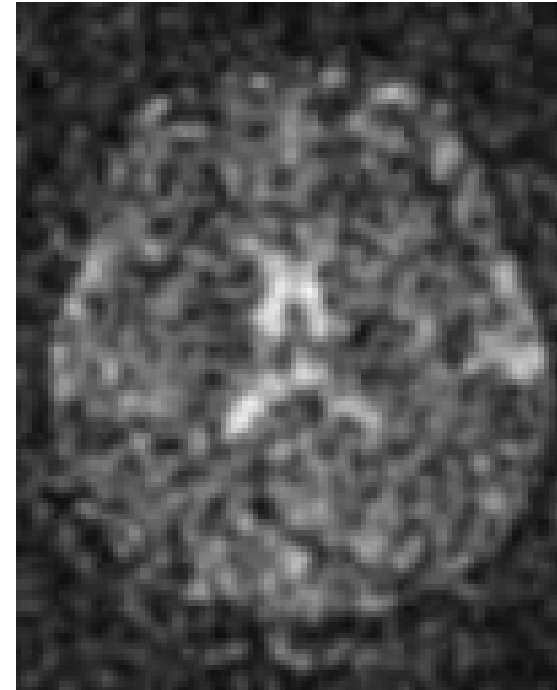
# First *In vivo* 9.4 T results



Anatomy - 1H  
MP-RAGE 4T  
1 mm isotropic  
5 min acq. time



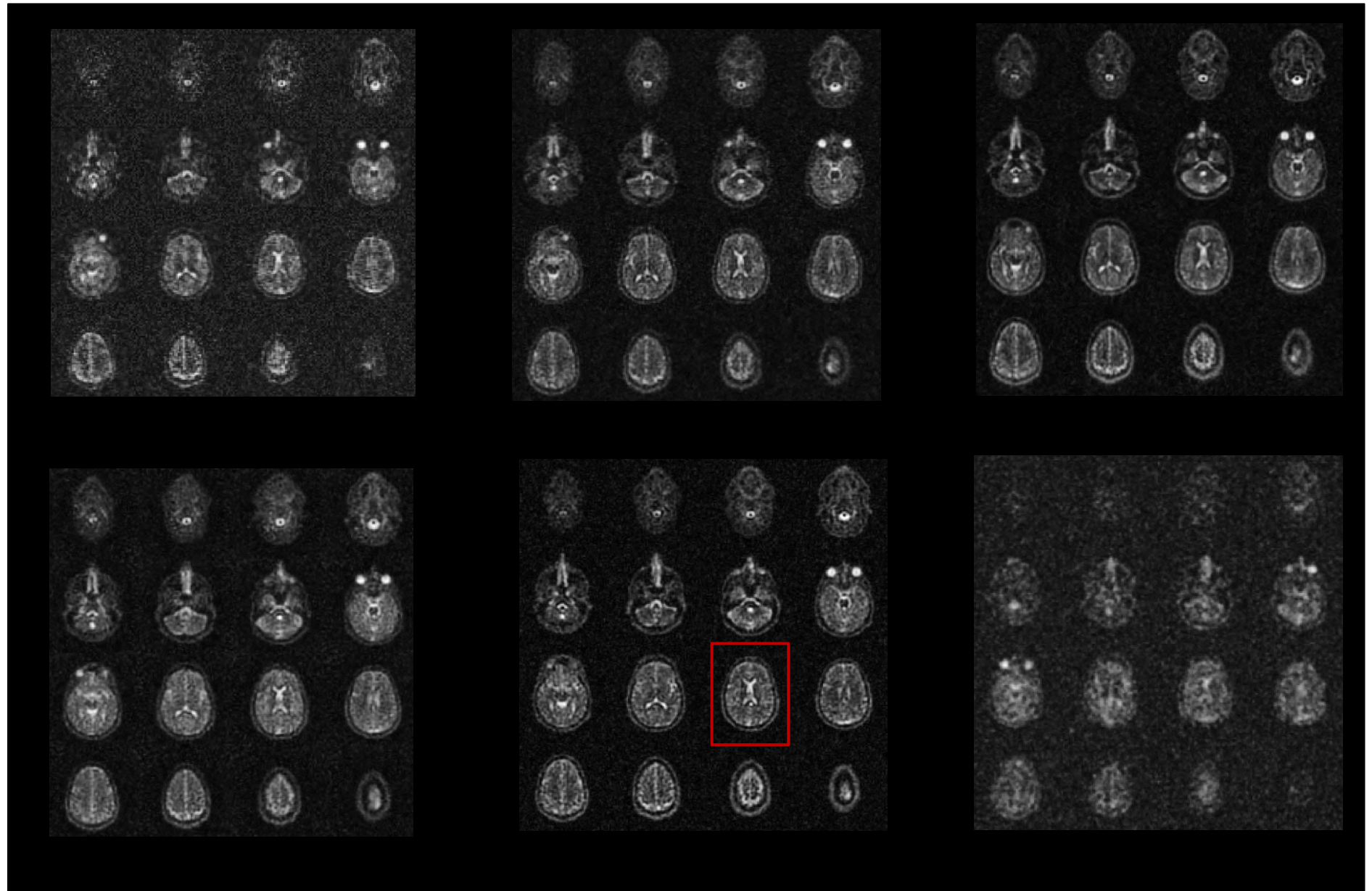
Sodium – 9.4T  
TPI  
2 mm isotropic  
15 min acq. time



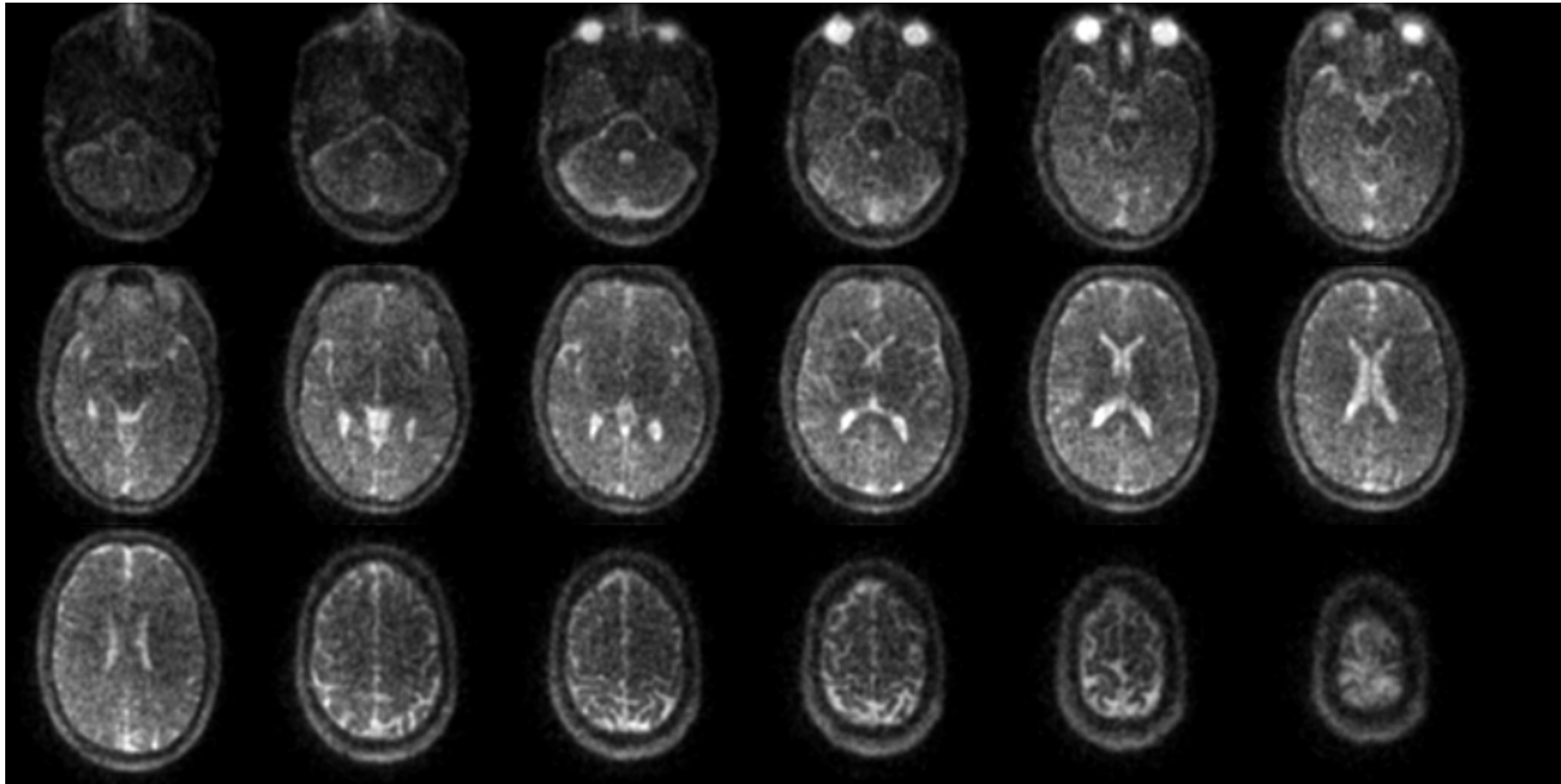
Sodium – 4T  
TPI  
2 mm isotropic  
15 min acq. time



# *In vivo* measurements ( $^{23}\text{Na}$ )



# Sodium Imaging ( $2 \times 2 \times 2 \text{mm}^3$ in 7 minutes)



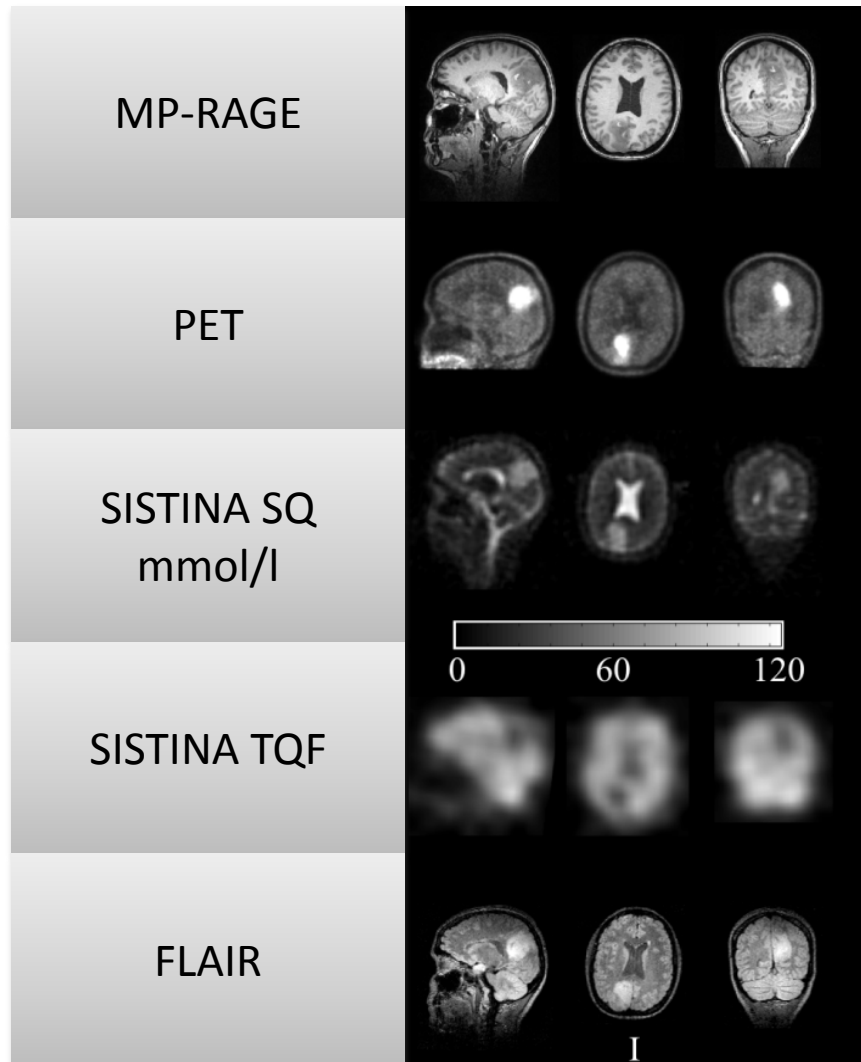
**In vivo  $^{23}\text{Na}$  Imaging with TPI steady state (TR/TE/flip = 50/0.4/60 [ms/ms/deg])**

Fiege, Romanzetti, ..., and Shah (2013) *Magn. Reson. Med.*

Fiege, Romanzetti, ..., and Shah (2013) *J. Magn. Reson.*

Romanzetti, ..., and Shah (2014) *NeuroImage (in press)*

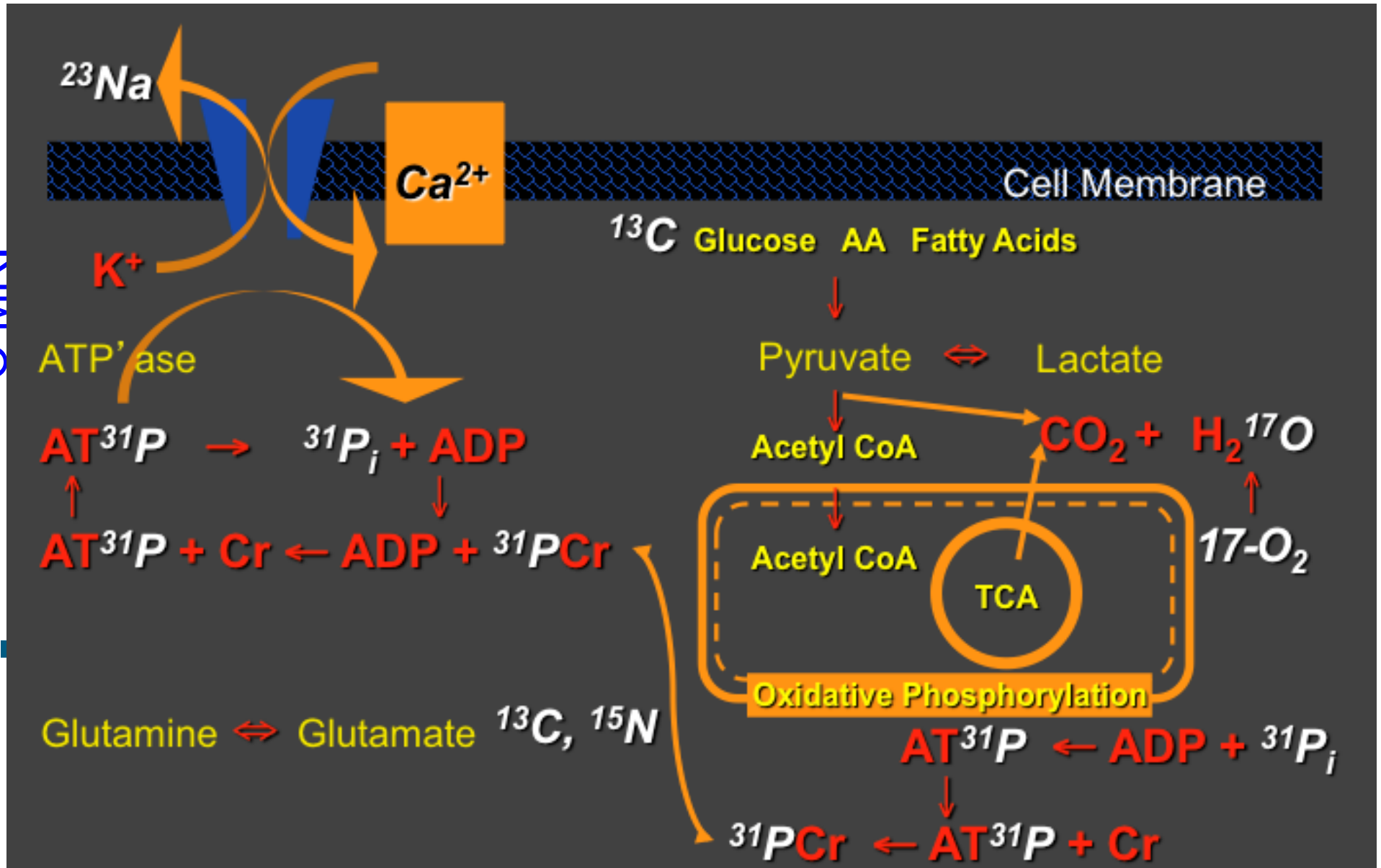
# $^{23}\text{Na}$ TQF *in vivo* 9.4 T results



*In vivo* results from three tumour patients.

# Goal: Non-invasive Quantitative Metabolic Imaging of Oxygen

<sup>17</sup>O MRI



# Oxygen Imaging

K. Moellenhoff and N. J. Shah

# $^1\text{H}$ MR Spectroscopy

D. Tse and N. J. Shah

# $^{31}\text{P}$ MR Spectroscopy

D. Tse and N. J. Shah

# Temporal Aspects

I. Neuner, J. Arrubla and N. J. Shah















S 300	S 300
6.7	7.7
2.2	2.8
S 300	S 300
1.7	7.6
2.8	2.2
S 200	S 200
6.7	7.7
1.1	1.2















SIEMENS

R+WASEL

Borgheln  
Düsseldor  
Horb, Vgl

0 22 71 465-0  
02 11 77 92 57-0  
0 33 1 47 46-0

H.C. WILSON

COLLASKIES

4 x 12M  
30 T

W003



BREUER & WASSER

ASEL  
REHKRANE

SIEMENS

SIEMENS

magnex  
scientific

BM BW 508



SIEMENS

magnex scientific

SIEMENS



SIEMENS

magnex  
scientific



SIEMENS



IMPORTANT  
BORN





# Acknowledgements

Dr. J. Felder

A. Celik

K. Vahedipour

C. Mirkes

MR Group

T. Okell / P. Jezzard (Oxford)

SIEMENS / BMBF

Dr. J. Scheins

Dr. E. Rota-Kops

L. Tellmann

PET Group

Dr. N. Galdiks

Dr. G. Stoffels

Dr. C. Filß

Brain Tumour Group

# Thank You!!

## Collaboration Opportunities

### Visiting Scholars / Guest Scientists

- Postdocs
- Graduate Students