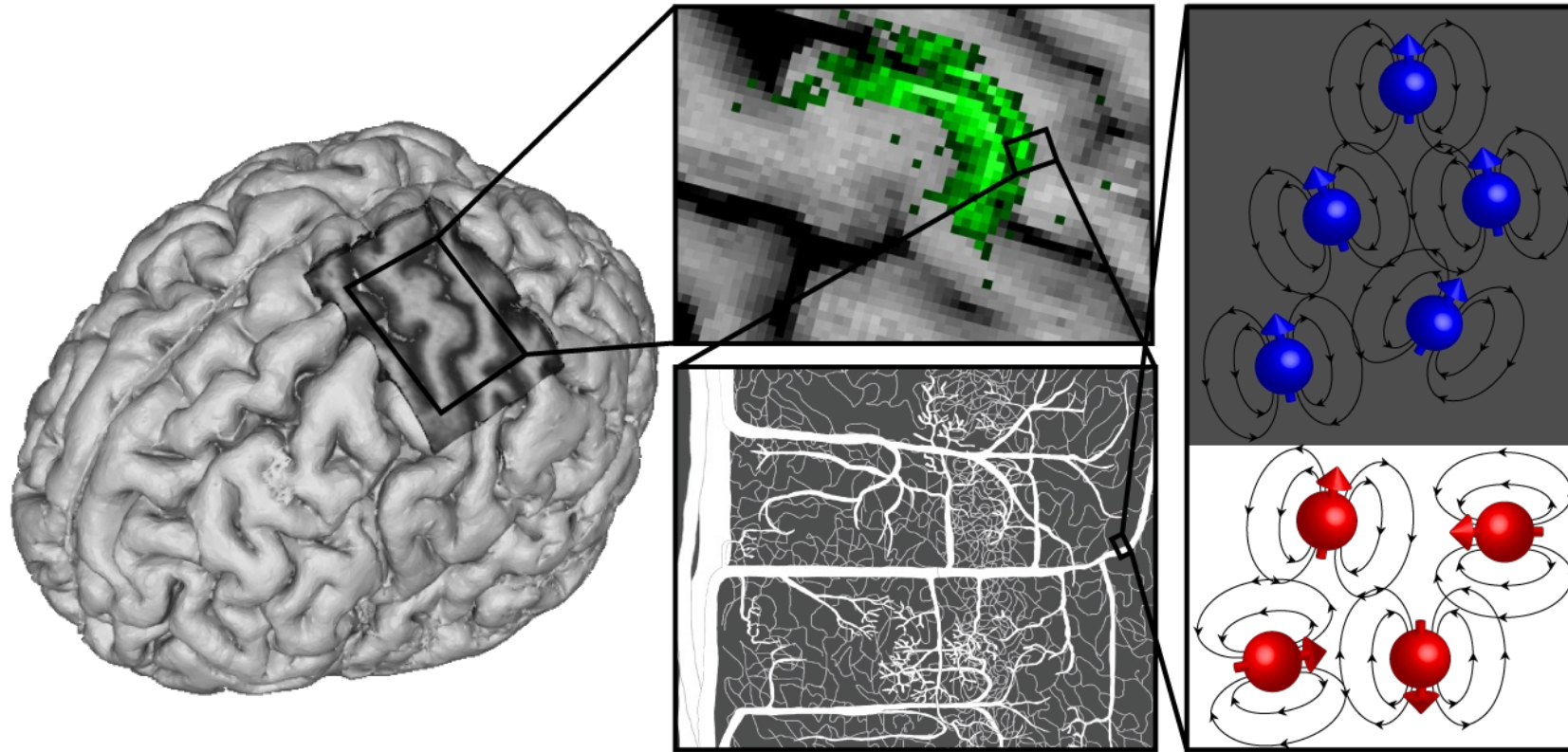


Functional contrast and processing strategies at high field and high resolution




Renzo (Laurentius) Huber



FMRI facility (FMRIF) at the NIMH/NINDS
of the National Institutes of Health (NIH)

Background Material

Relevant review article:



NeuroImage
Available online 20 July 2017
In Press, Corrected Proof



Non-BOLD contrast for laminar fMRI in humans: CBF, CBV, and CMRO₂

Laurentius Huber^a, Kamil Uludağ^{b,1}, Harald E. Möller^{c,1}

[Show more](#)

<https://doi.org/10.1016/j.neuroimage.2017.07.041> [Get rights and content](#)

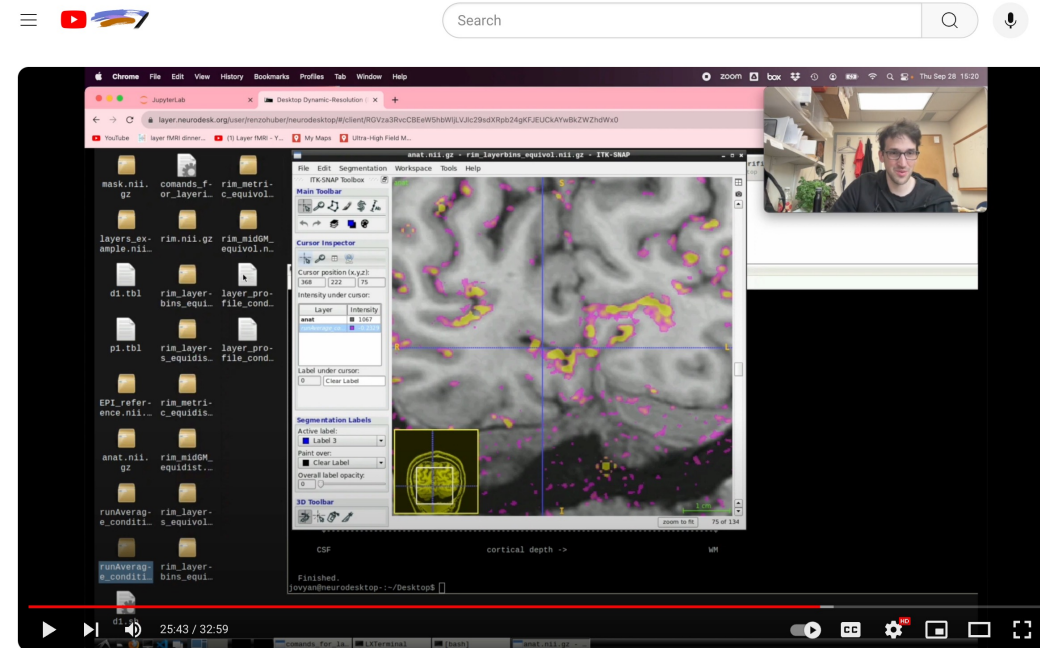
Highlights

- Non-BOLD-fMRI methods are reviewed for layer-dependent application in humans.
- ASL, VASO, and calibrated fMRI are promising candidates.
- Representative depth-dependent ASL/VASO/calibrated fMRI results are shown.
- ASL/VASO/calibrated fMRI are less sensitive but more specific than GE-BOLD.

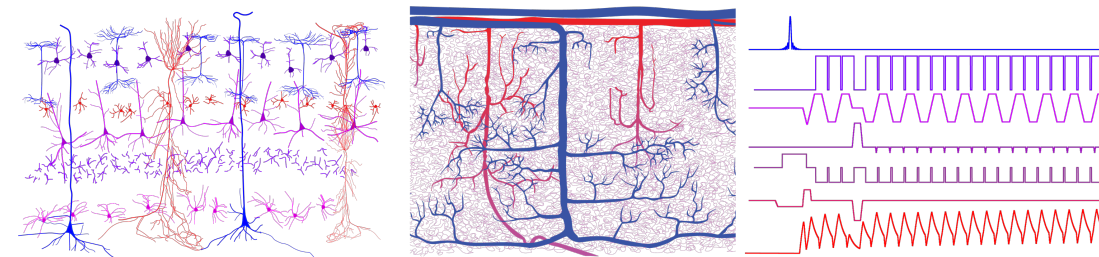
Abstract

Functional magnetic resonance imaging (fMRI) using the blood oxygenation level-dependent (BOLD) contrast indirectly probes neuronal activity changes via evoked

Hand-on layerification, https://youtu.be/tSA77mFTwcg?si=cQ_PqFXFu-VBINxX

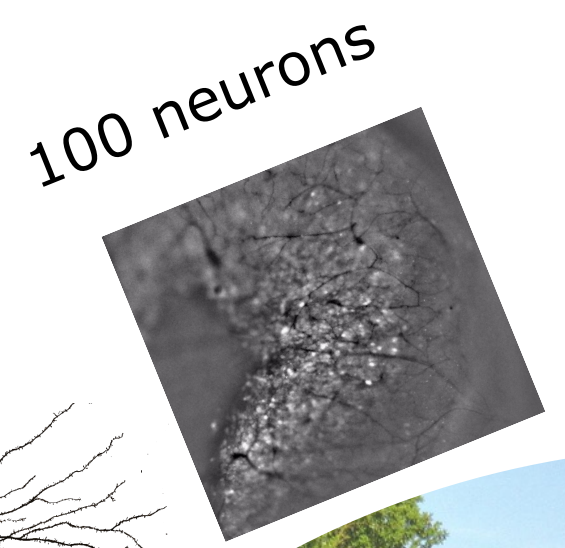
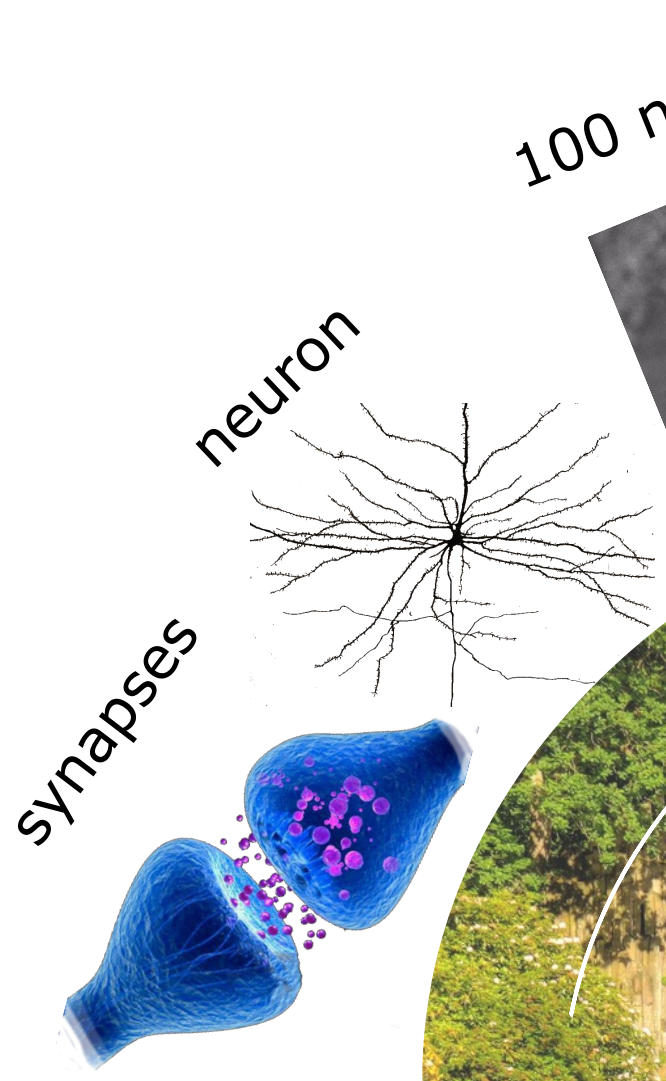


<https://education.martinos.org/workshop-on-laminar-fmri/>

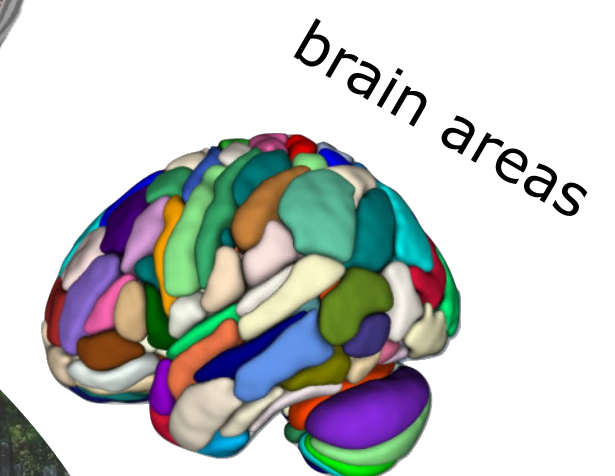
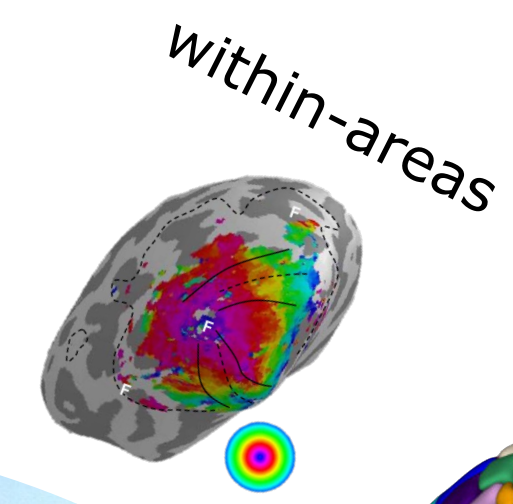
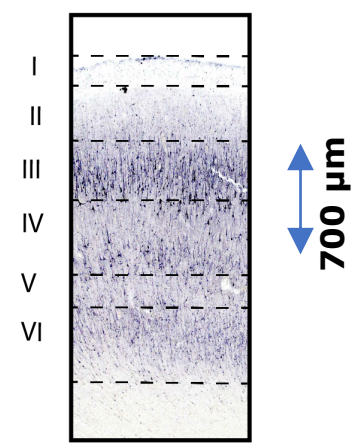


Ask me for recordings

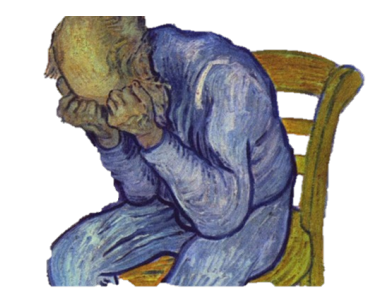
Why high resolutions?



layers & columns



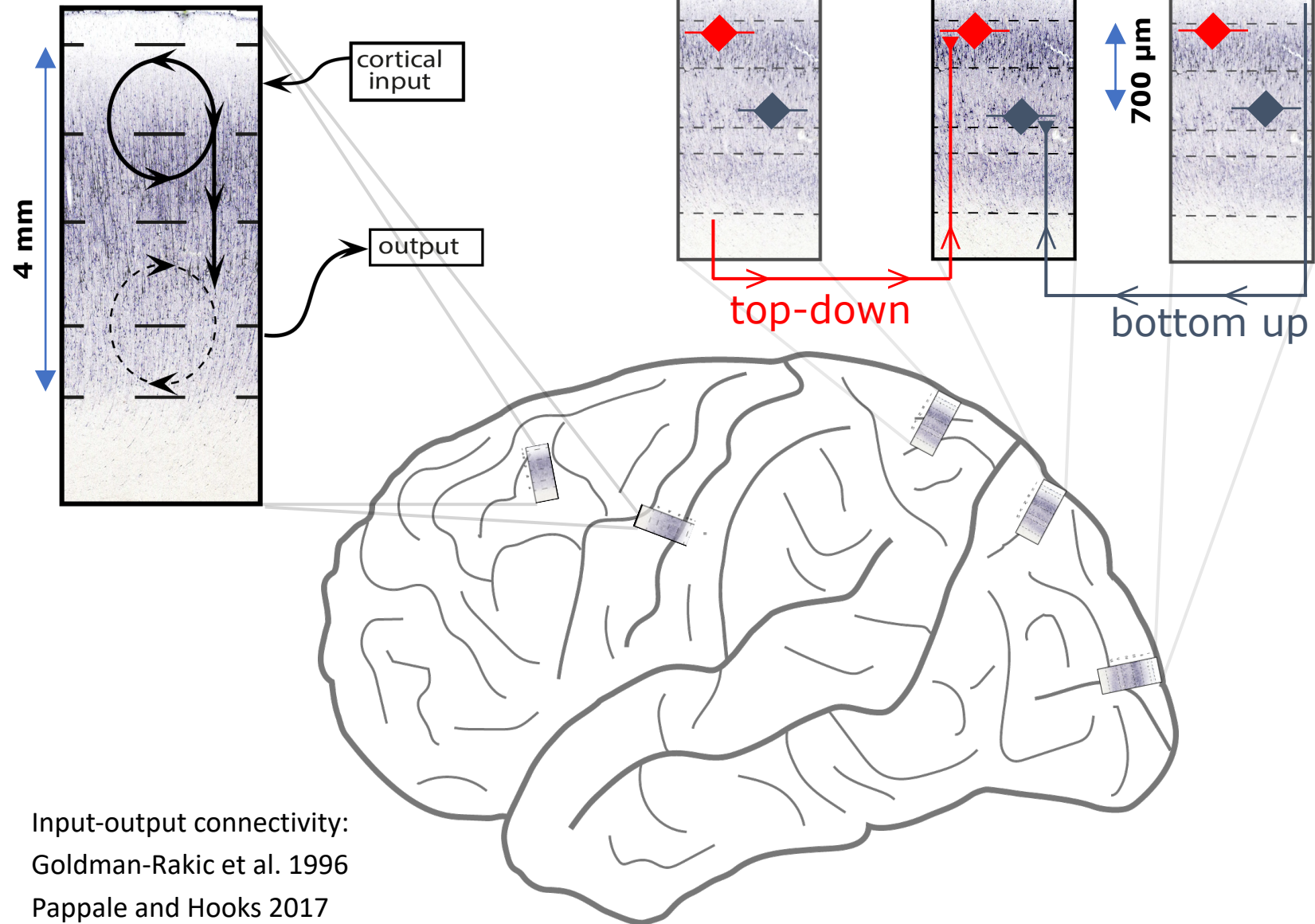
behavior



Layers provide directionality information

higher hierarchy

lower hierarchy

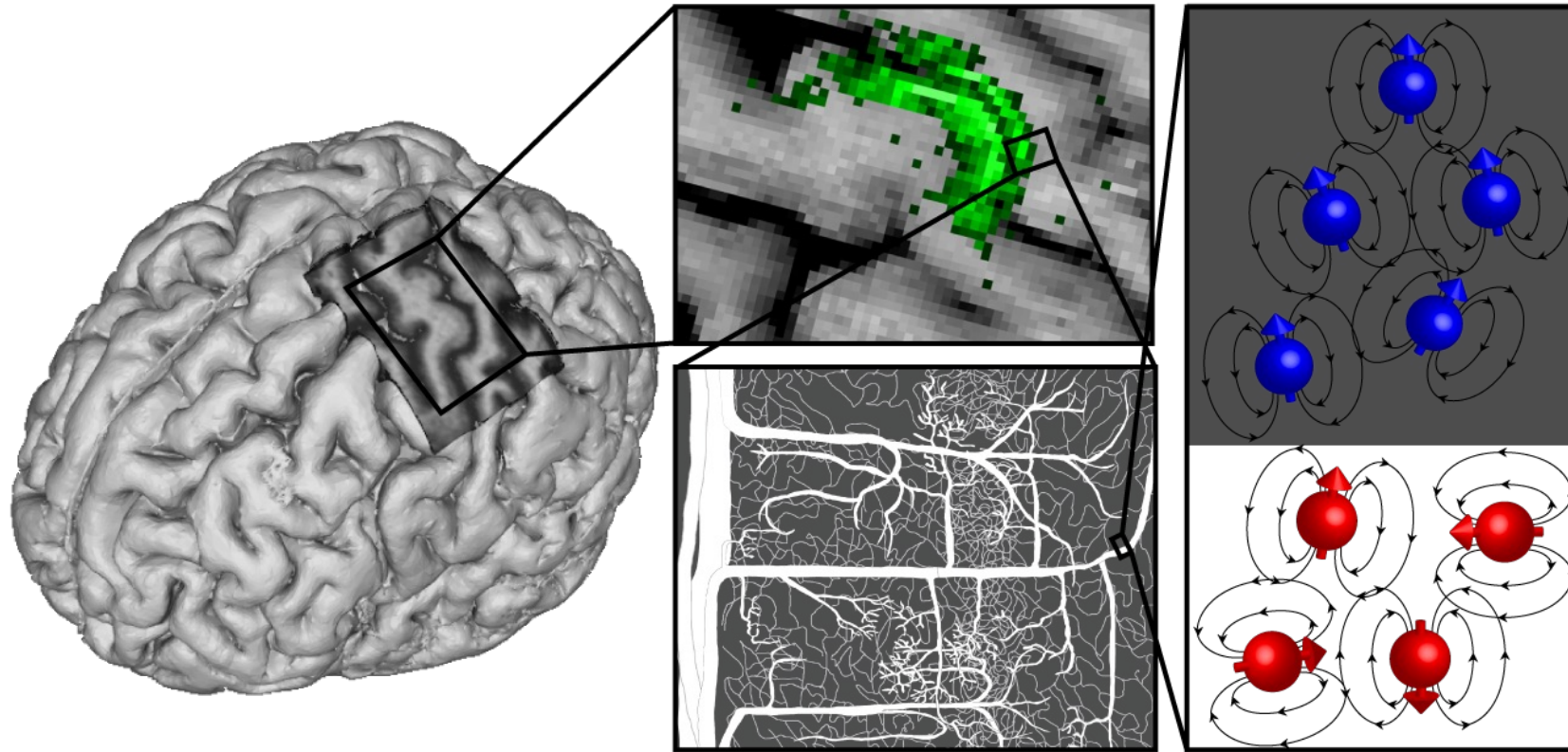


Hierarchical connectivity in the canonical micro-circuit:

Felleman and Van Essen 1991

Input-output connectivity:
Goldman-Rakic et al. 1996
Pappale and Hooks 2017

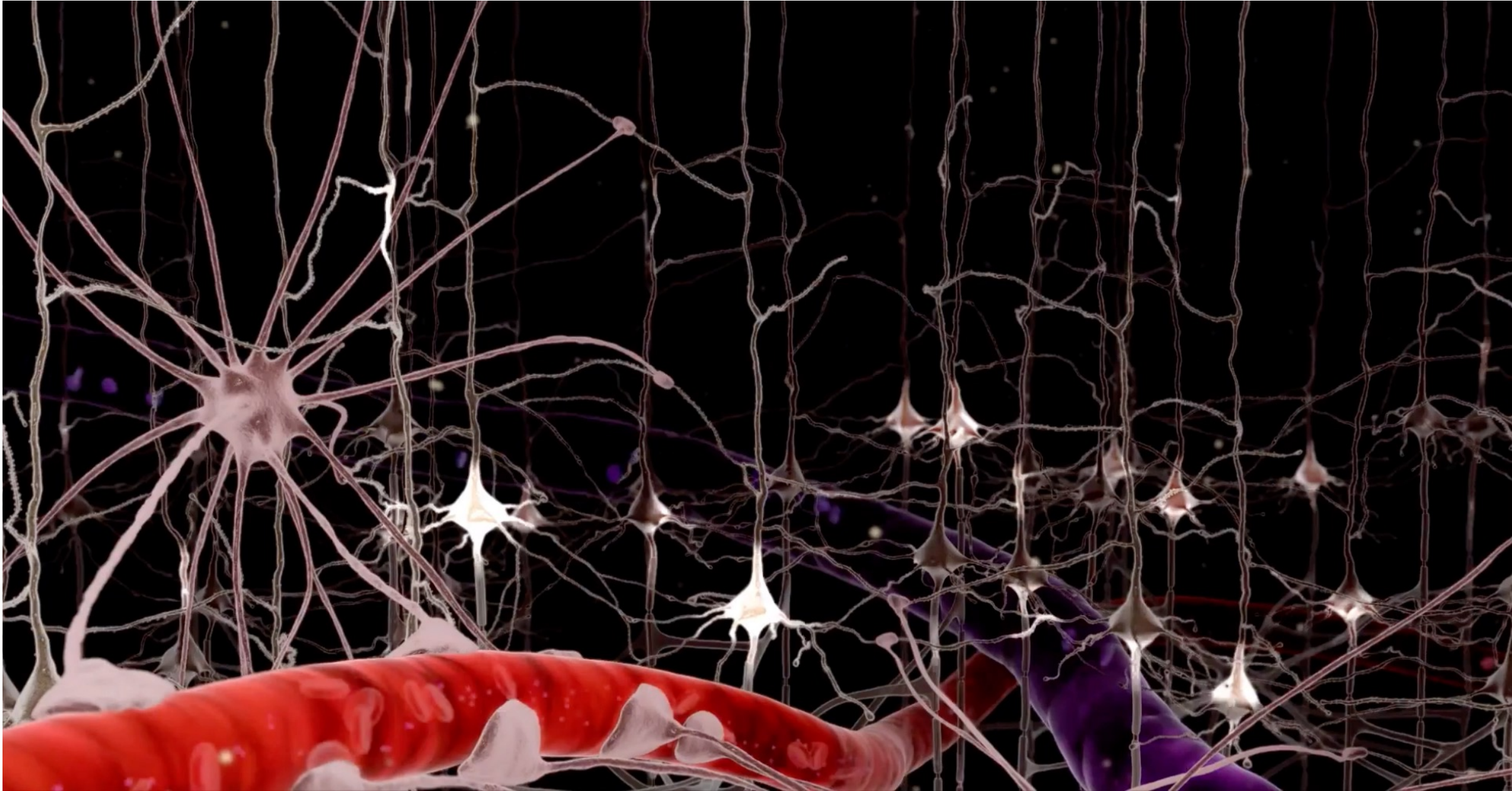
Functional contrast and processing strategies at high field and high resolution



Renzo (Laurentius) Huber

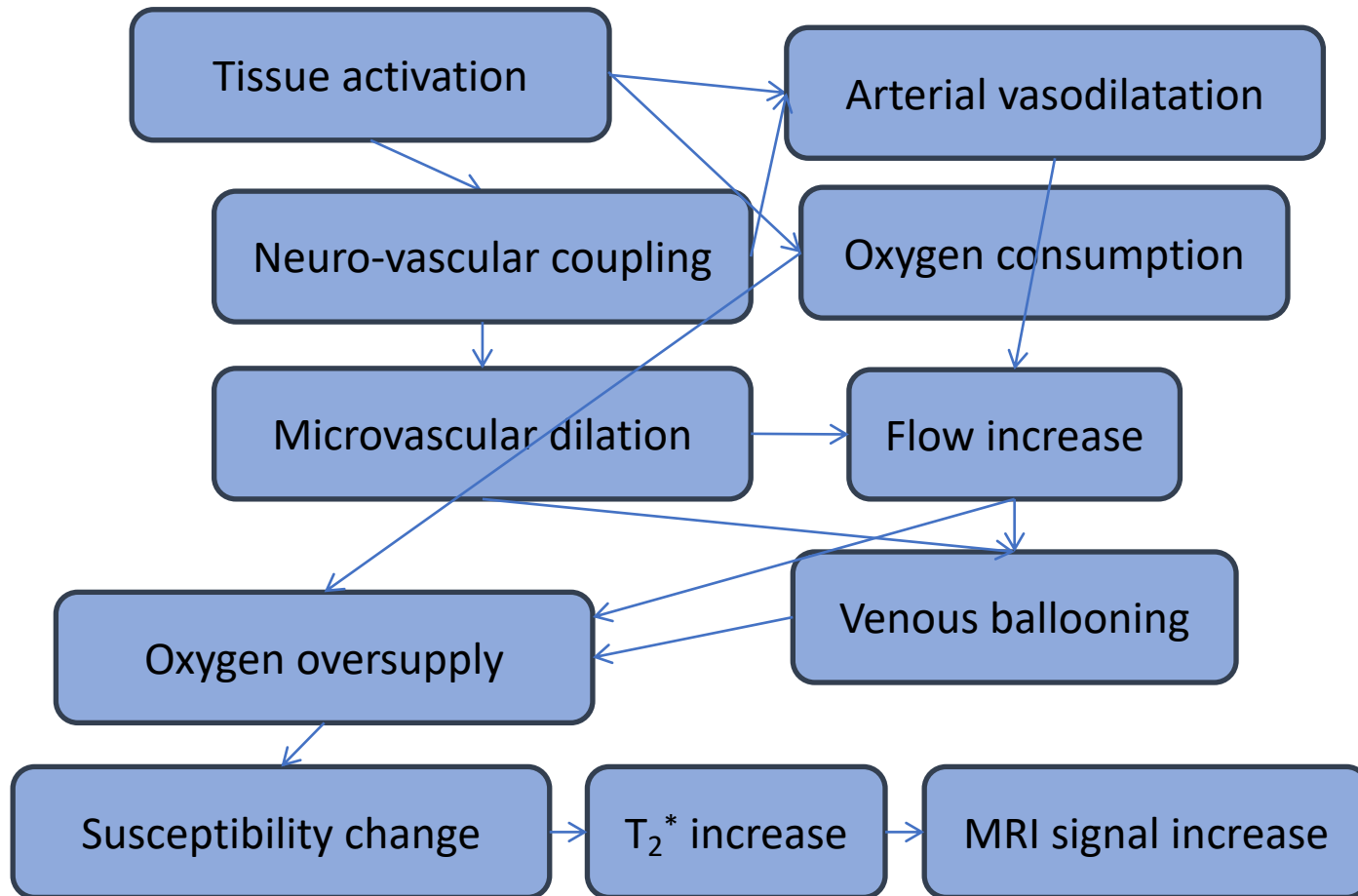
FMRI facility (FMRIF) at the NIMH/NINDS
of the National Institutes of Health (NIH)

Neurovascular coupling: fMRI is an indirect measure of neural activity



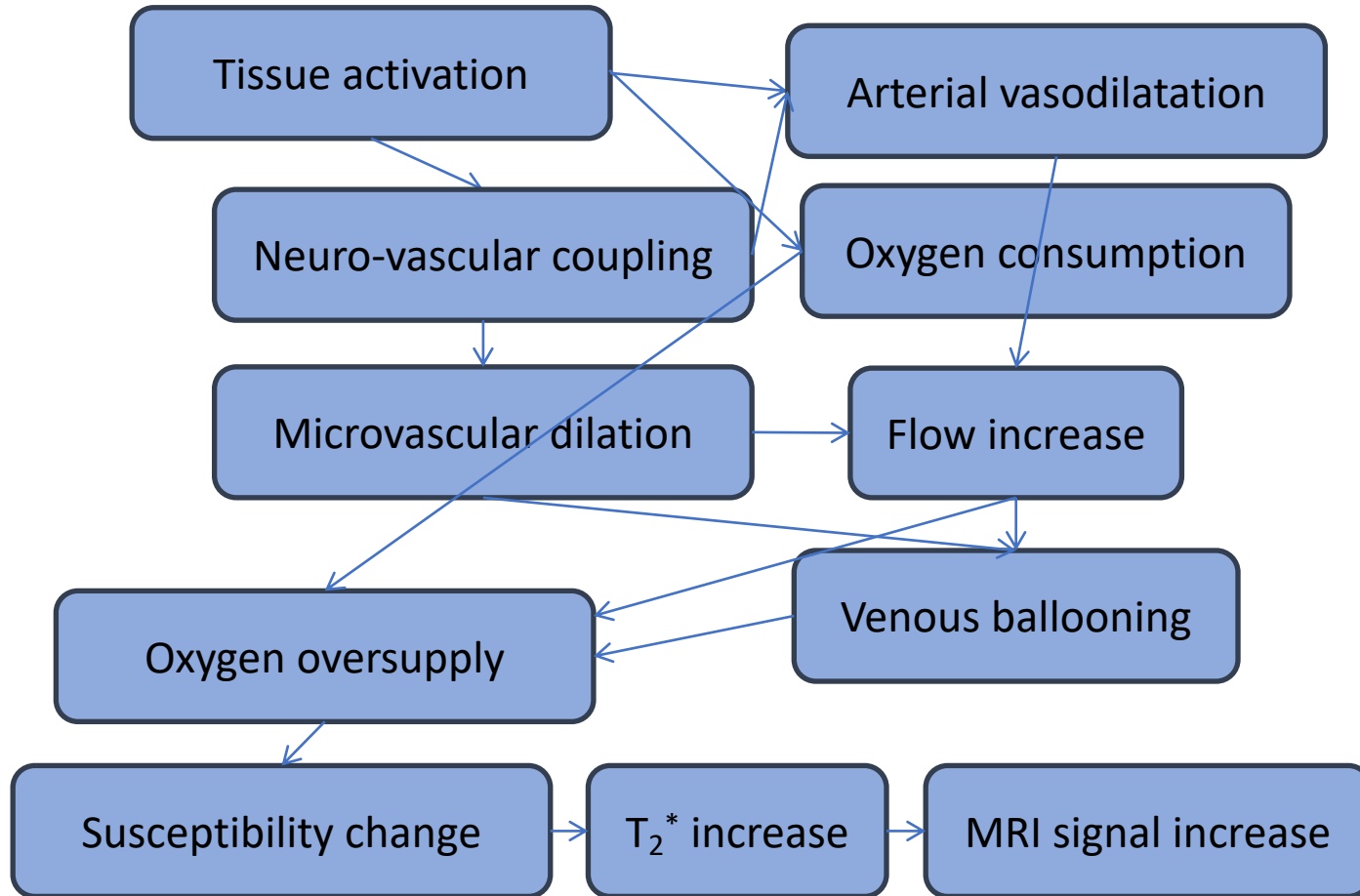
Quantifiability of GE-BOLD

BOLD contrast origin is not straight-forward:

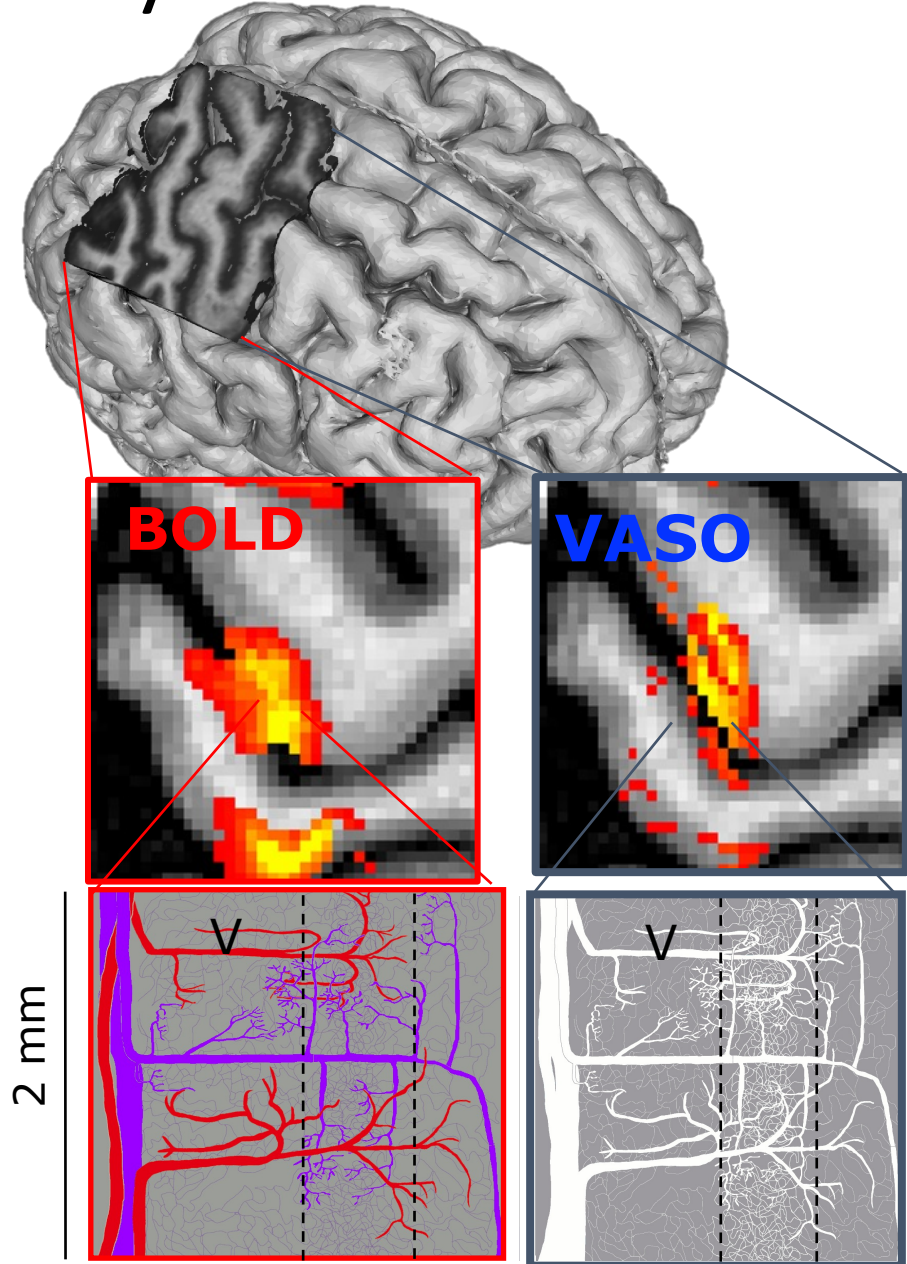


Quantifiability of GE-BOLD

BOLD contrast origin is not straight-forward:



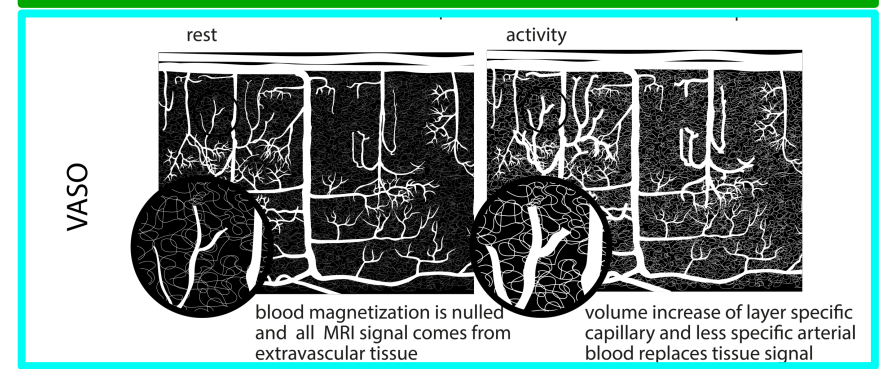
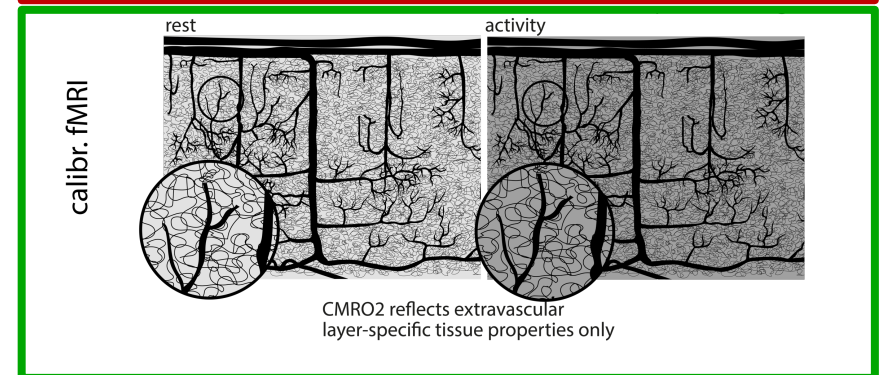
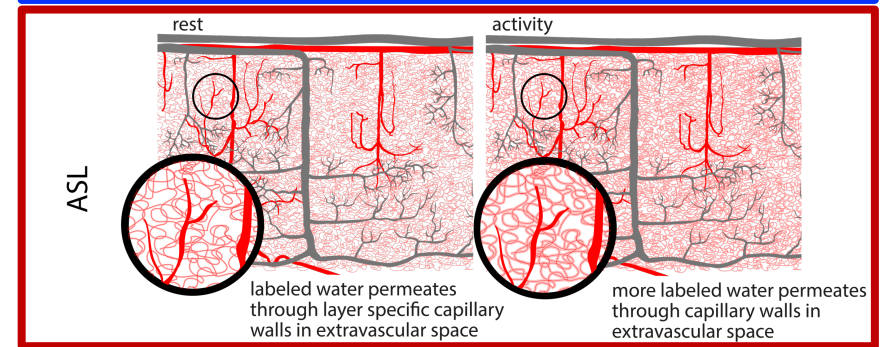
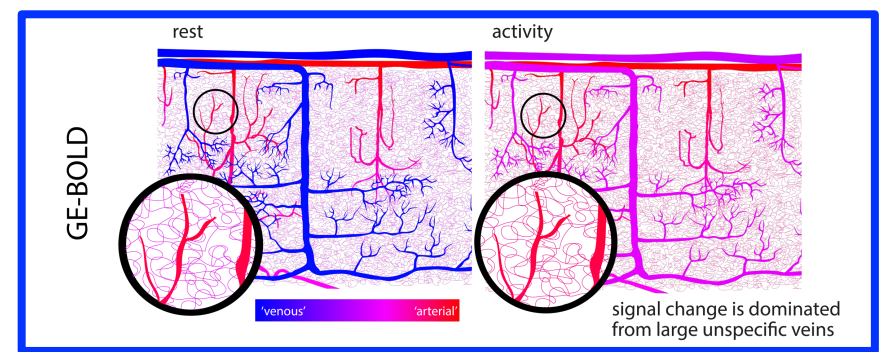
Specificity of GE-BOLD



[Huber et al., NeuroImage, 2015]

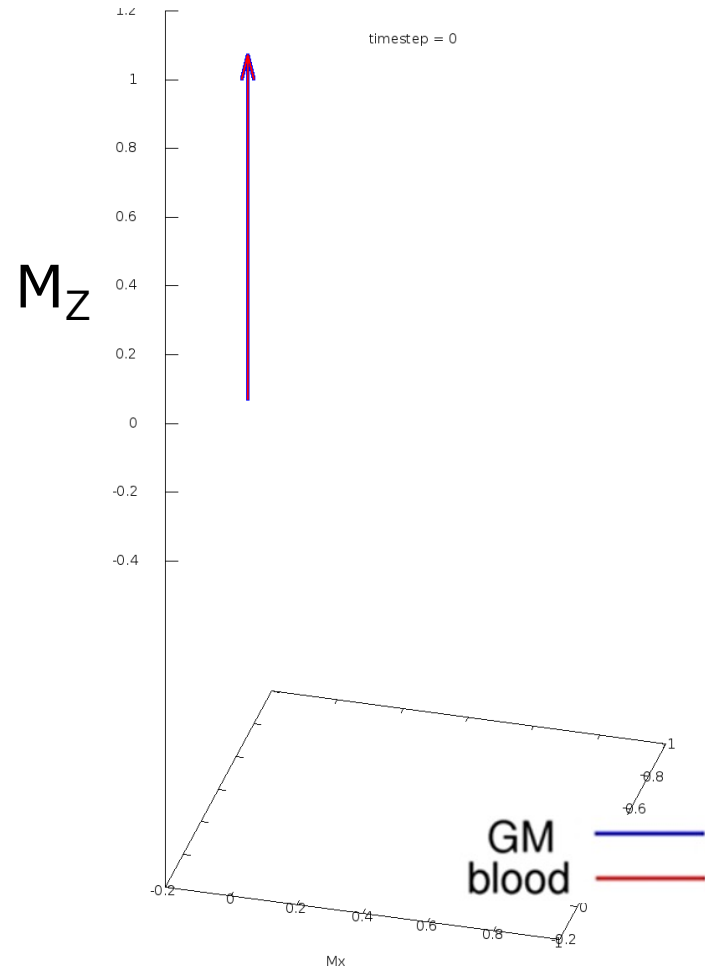
[Lu et al., MRM, 2003]

Popular alternative contrast



CBV contrast in Vascular Space Occupancy (VASO)

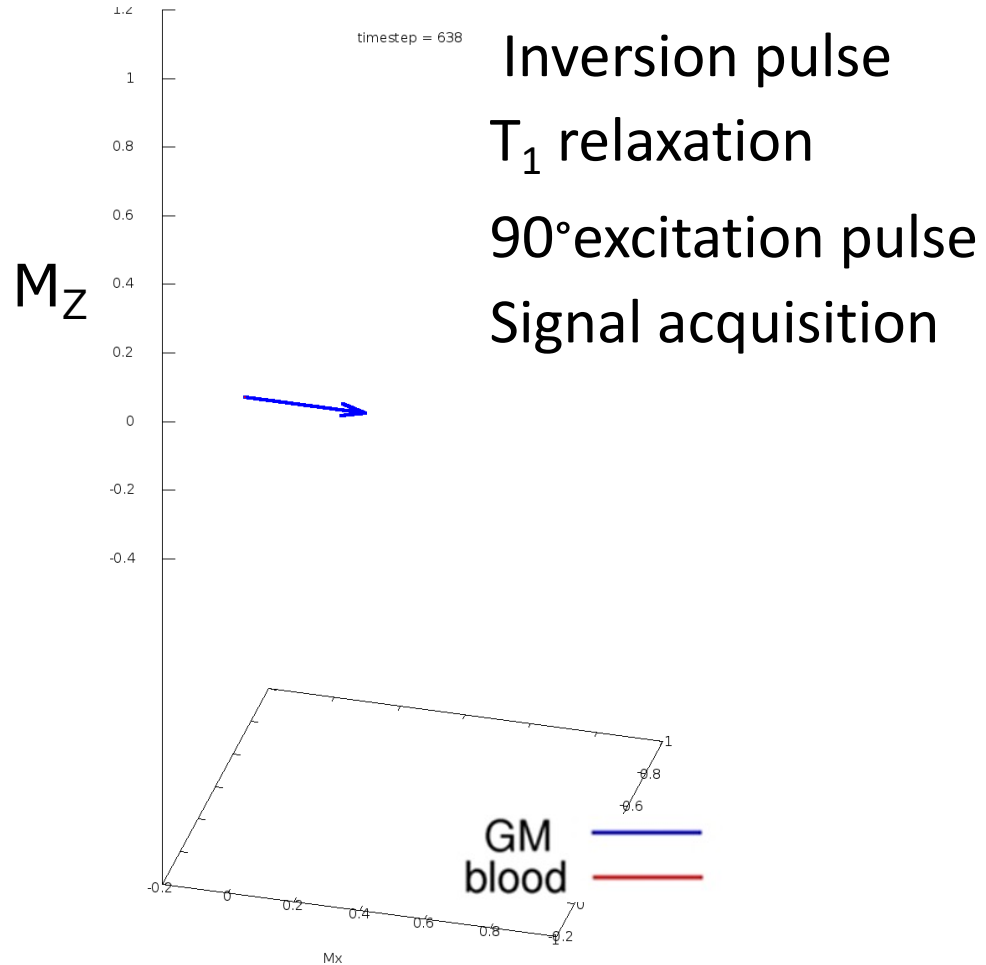
VASO is based on blood nulling in an inversion-recovery sequence



[Lu et al., MRM, 2003; Jin and Kim 2008]

CBV contrast in Vascular Space Occupancy (VASO)

VASO is based on blood nulling in an inversion-recovery sequence



Inversion pulse

T_1 relaxation

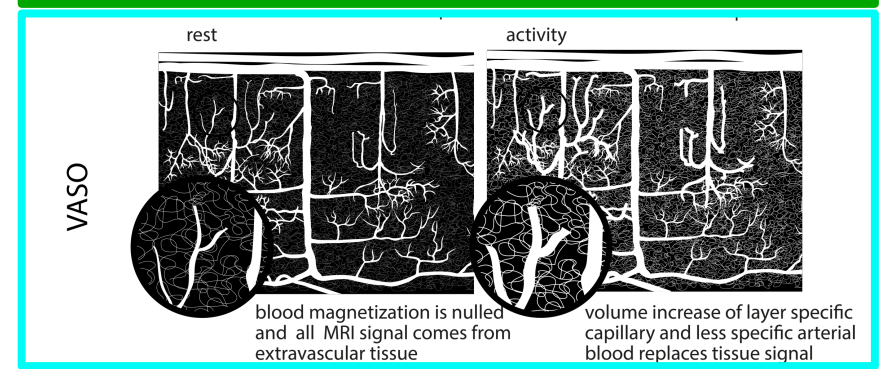
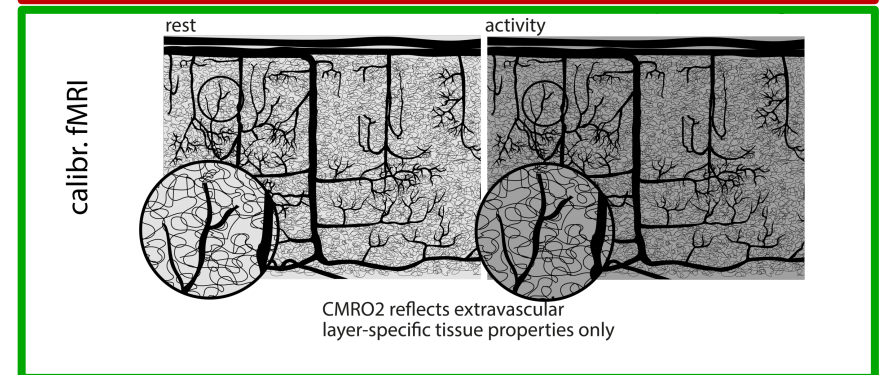
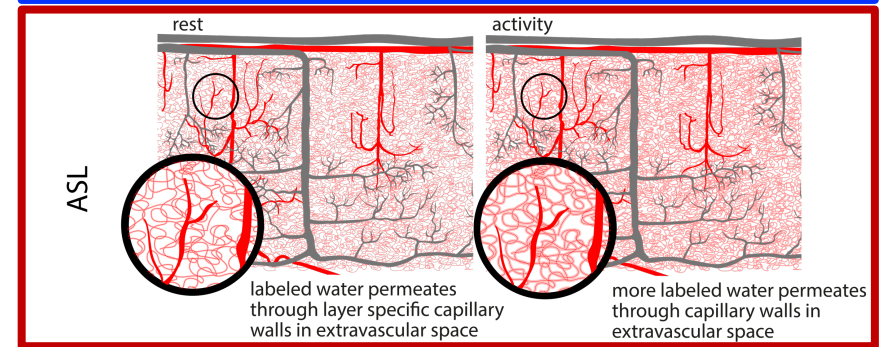
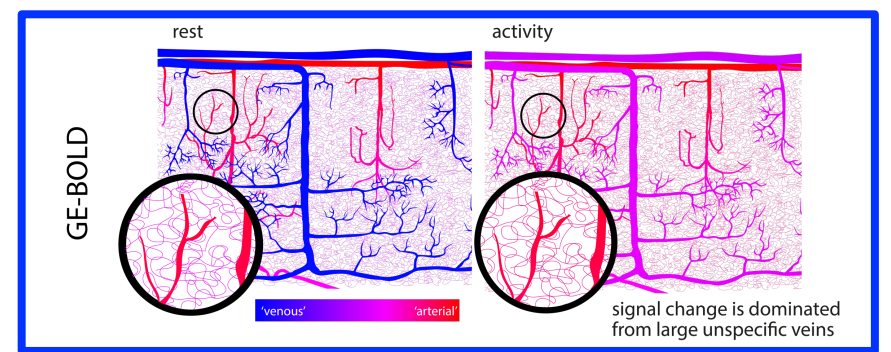
90° excitation pulse

Signal acquisition

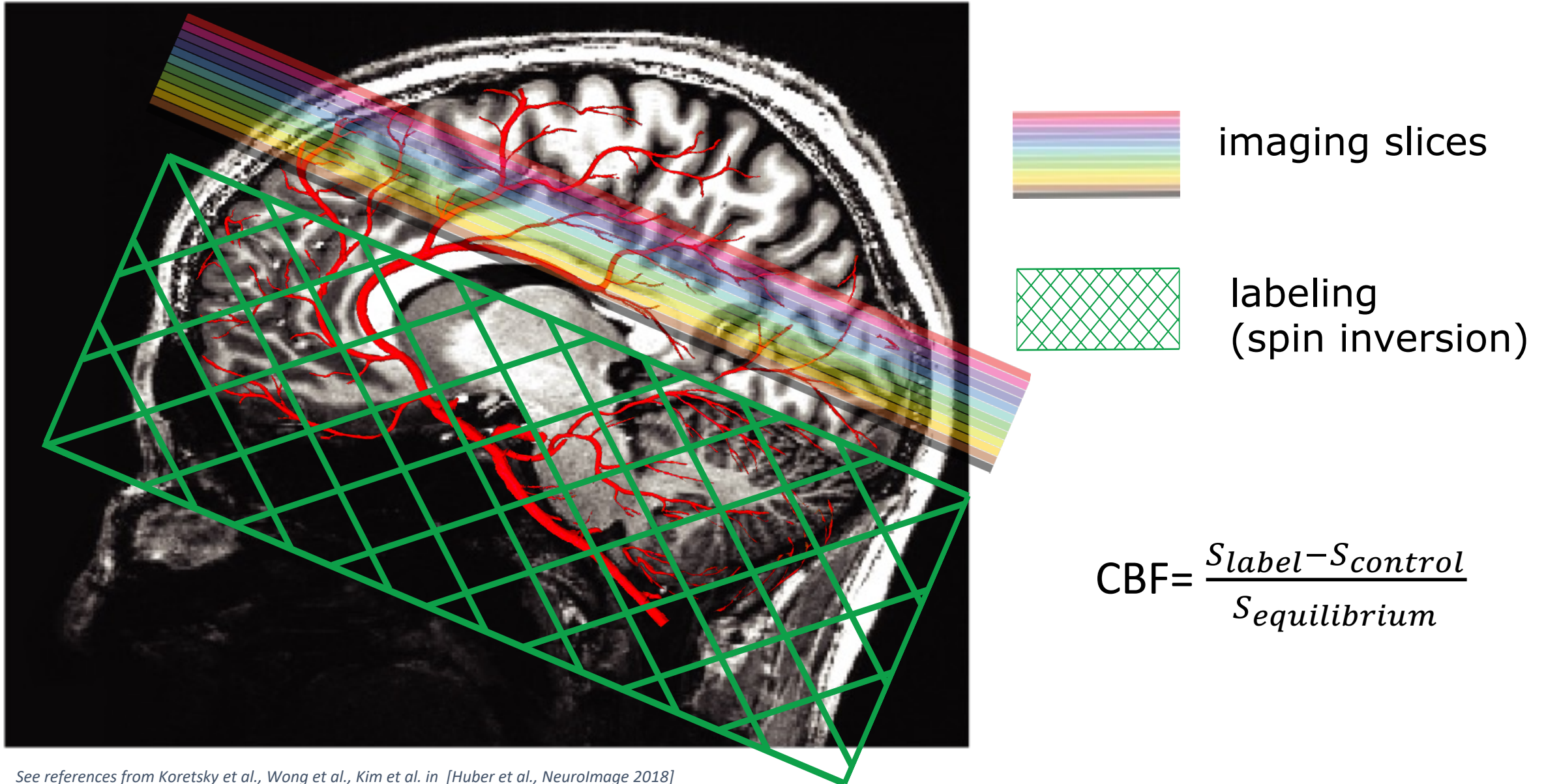
VASO is a negative contrast

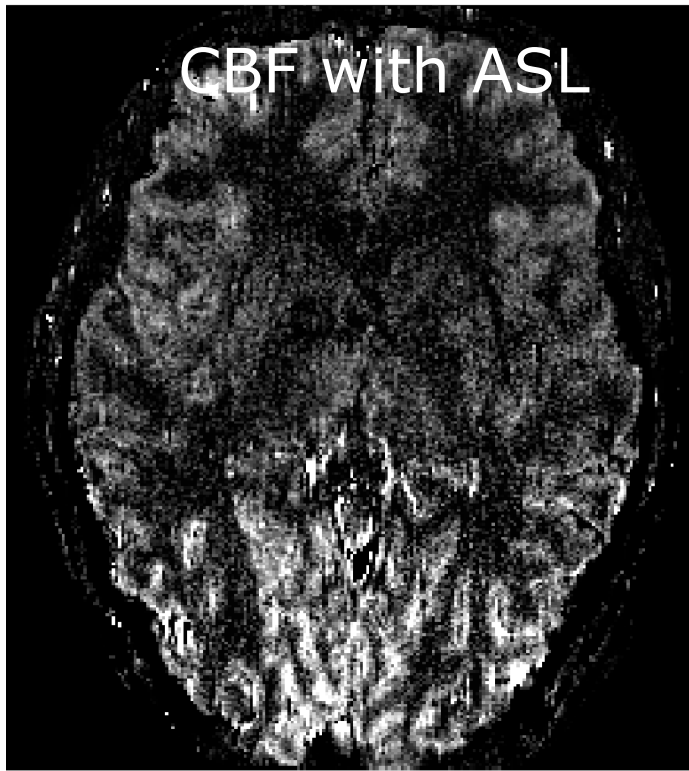


Popular alternative contrast

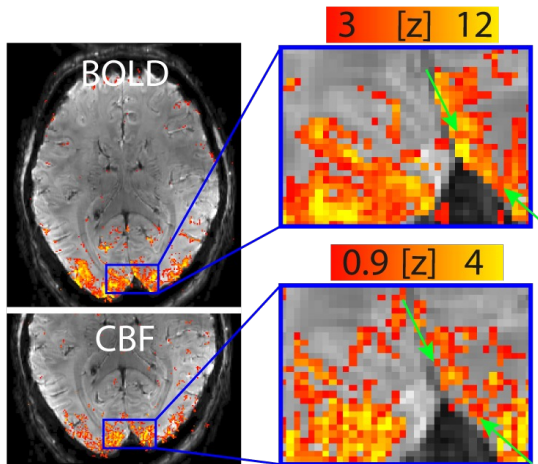
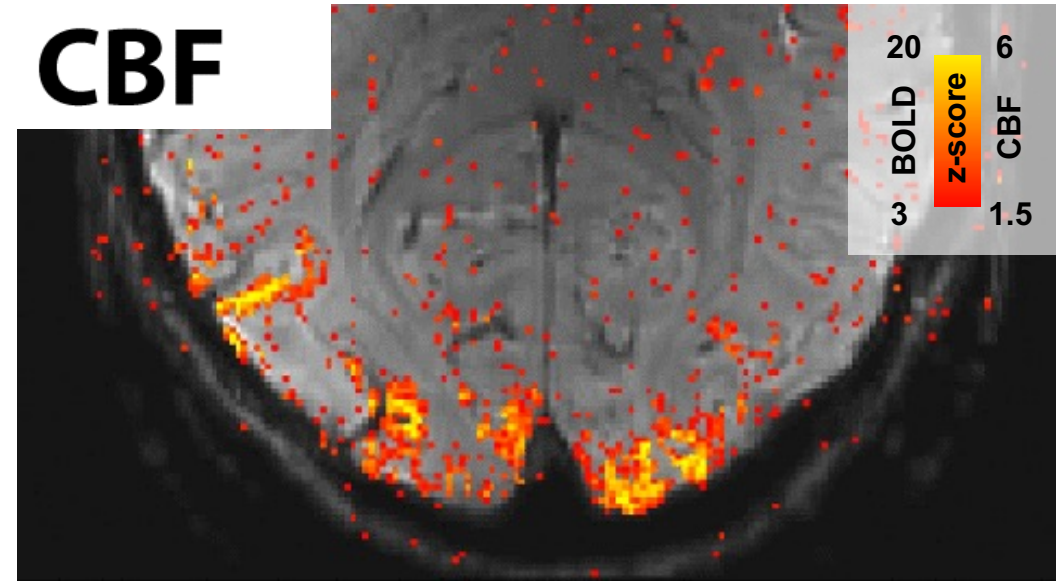


ASL (arterial spin labeling) for CBF measurements (FAIR)

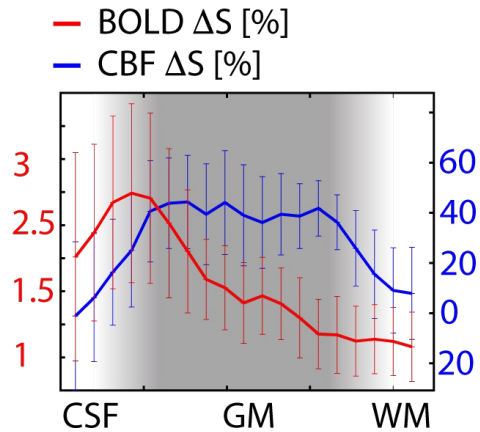




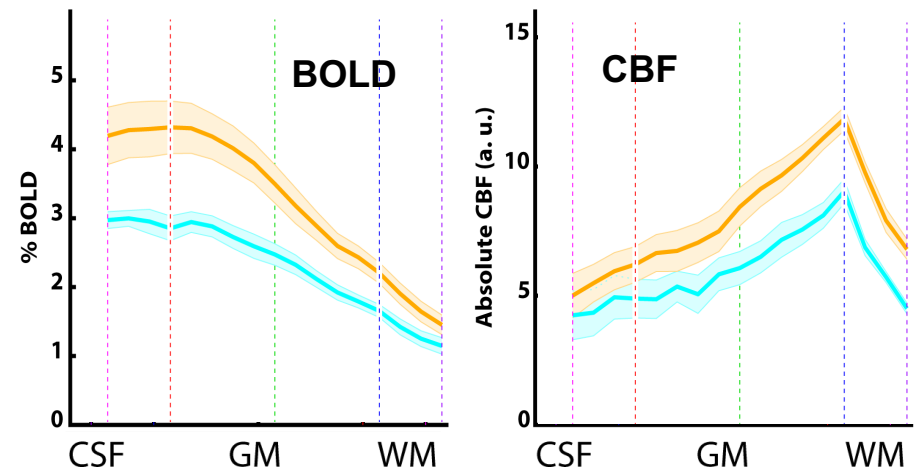
functional CBF with ASL



resolution: 0.8 mm



0.9mm iso, FAIR PASL, 7T

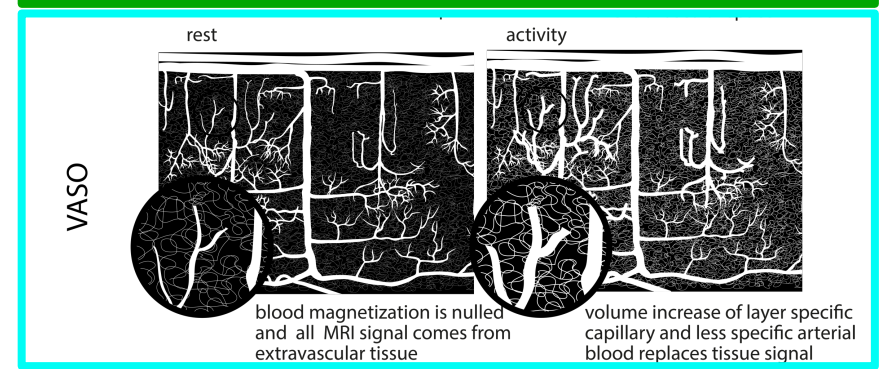
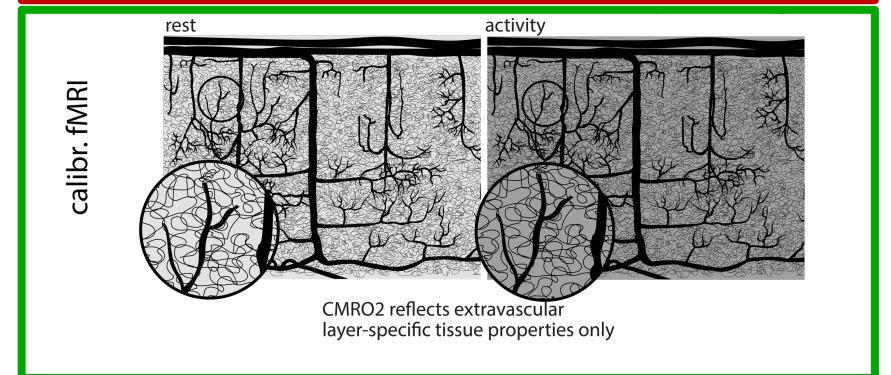
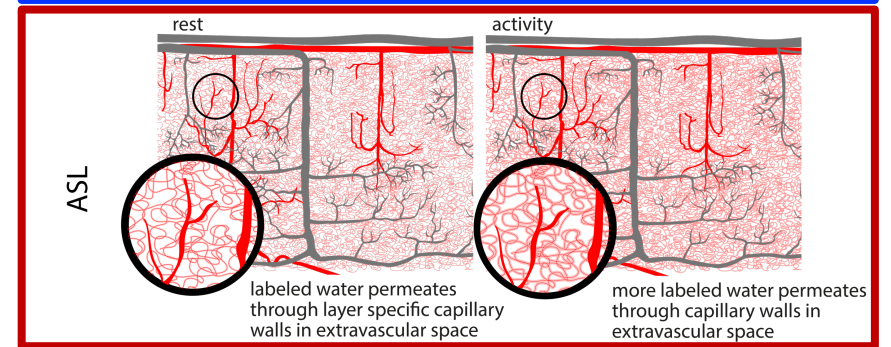
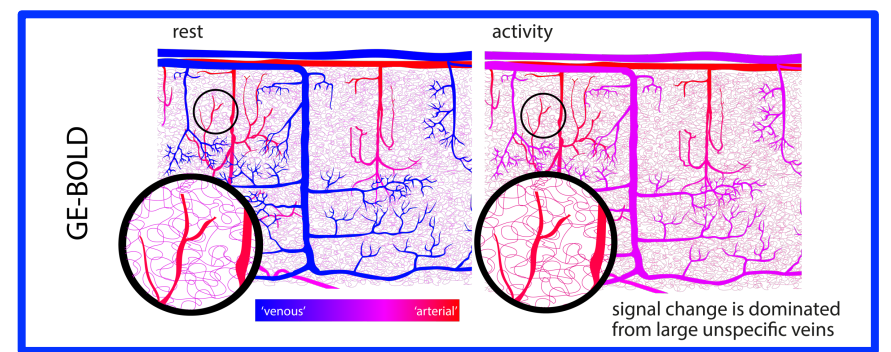


adapted from [Kashyap, PLOS One, 2021]

Data from Dimo Ivanov, MBIC, Maastricht

[Huber, Uludağ, Möller, NeuroImage, 2018]

Popular alternative contrast



Calibrated BOLD: estimation of CMRO₂

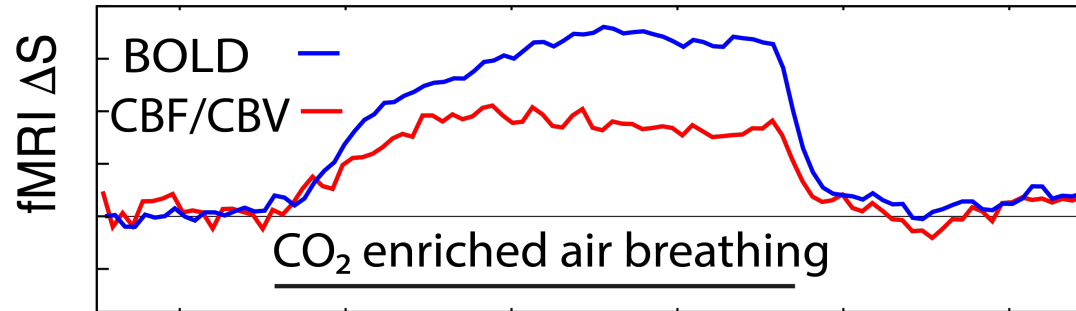
$$\frac{\Delta BOLD}{BOLD} = M \left[1 - \left(\frac{CMR_{O_2}}{CMR_{O_2}|_{rest}} \right)^\beta \left(\frac{CBV_v}{CBV_v|_{rest}} \right)^{\frac{\alpha_v - \beta}{\alpha_t}} \right]$$

[Hoge, 1999], [Davis, 1998]

hypercapnia

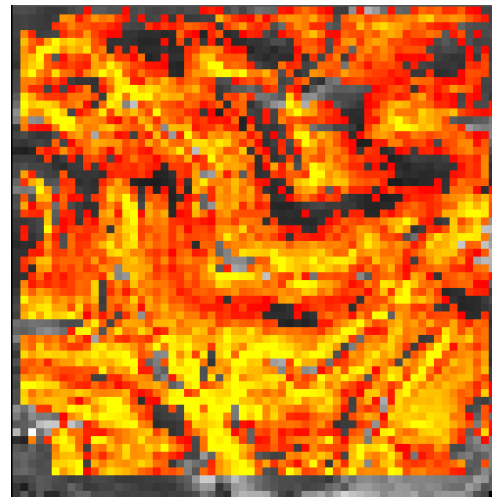
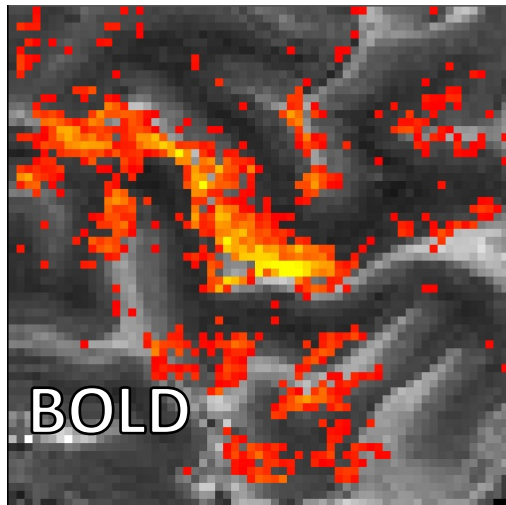
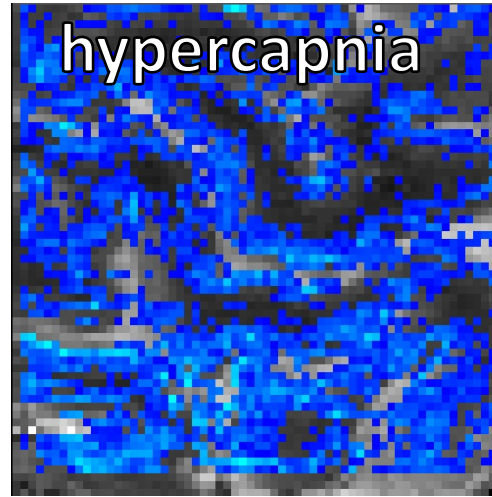
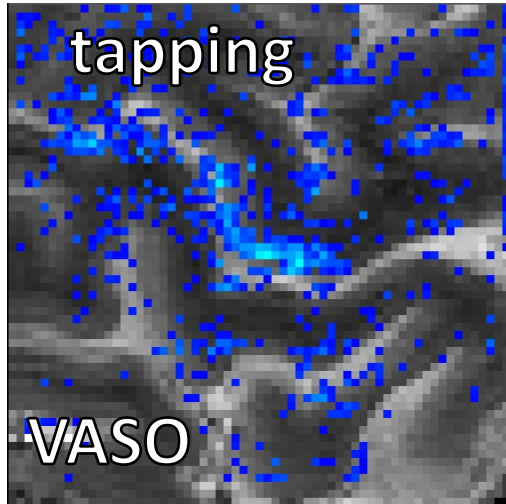
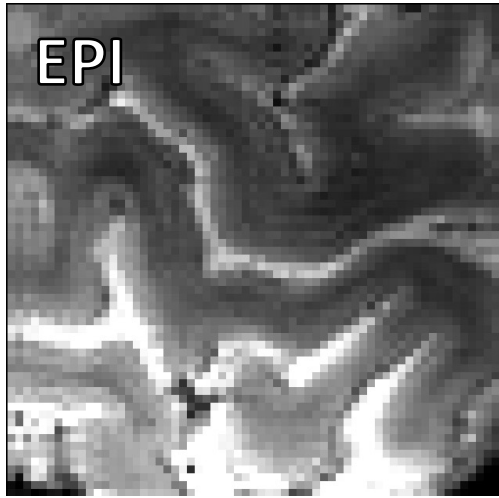
	Vessel dilation	Oxygen consumption
gas calibration	✓	✗

calibration experiment

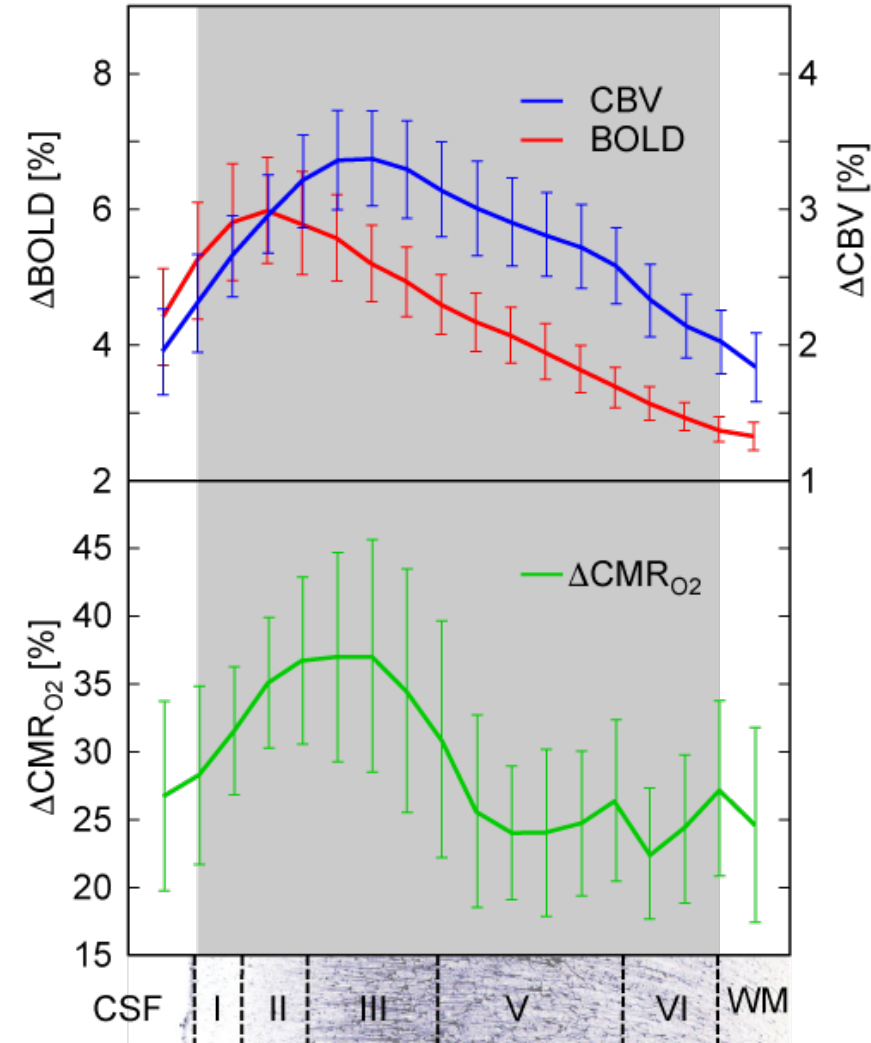


Calibrated BOLD: estimation of CMRO₂

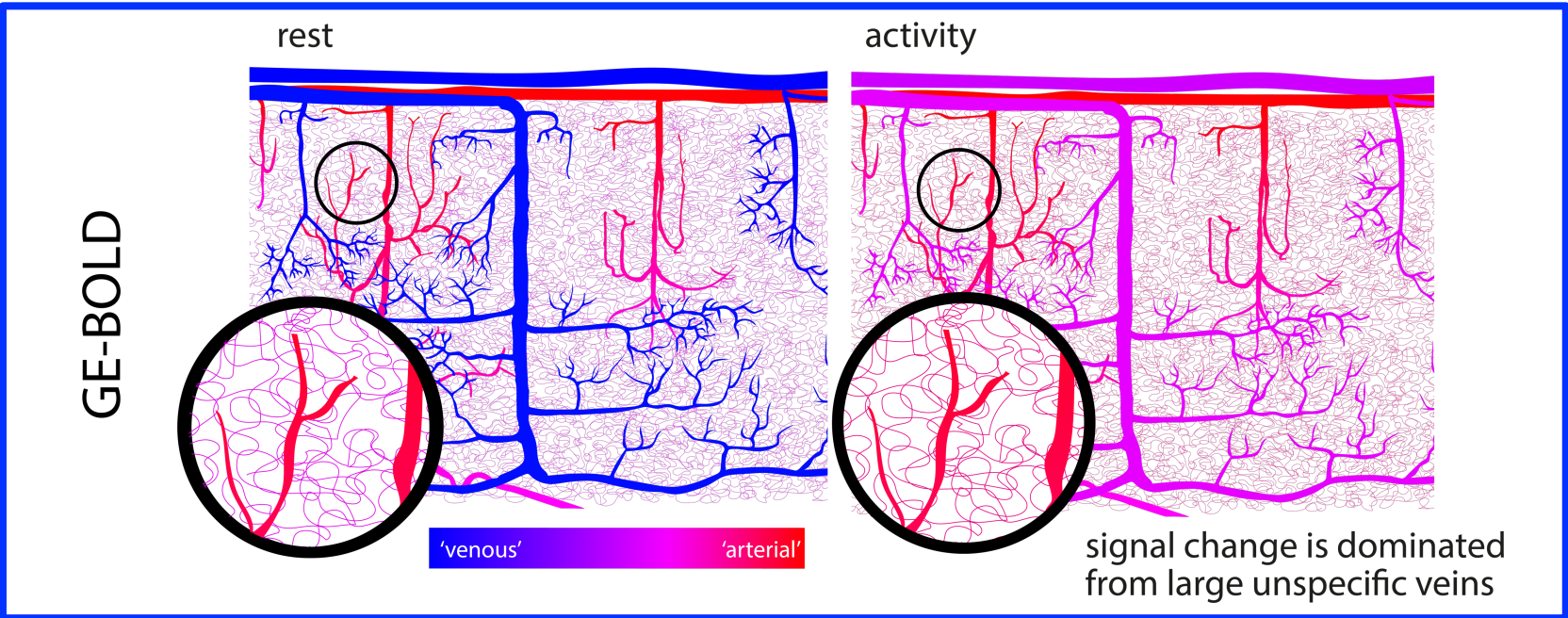
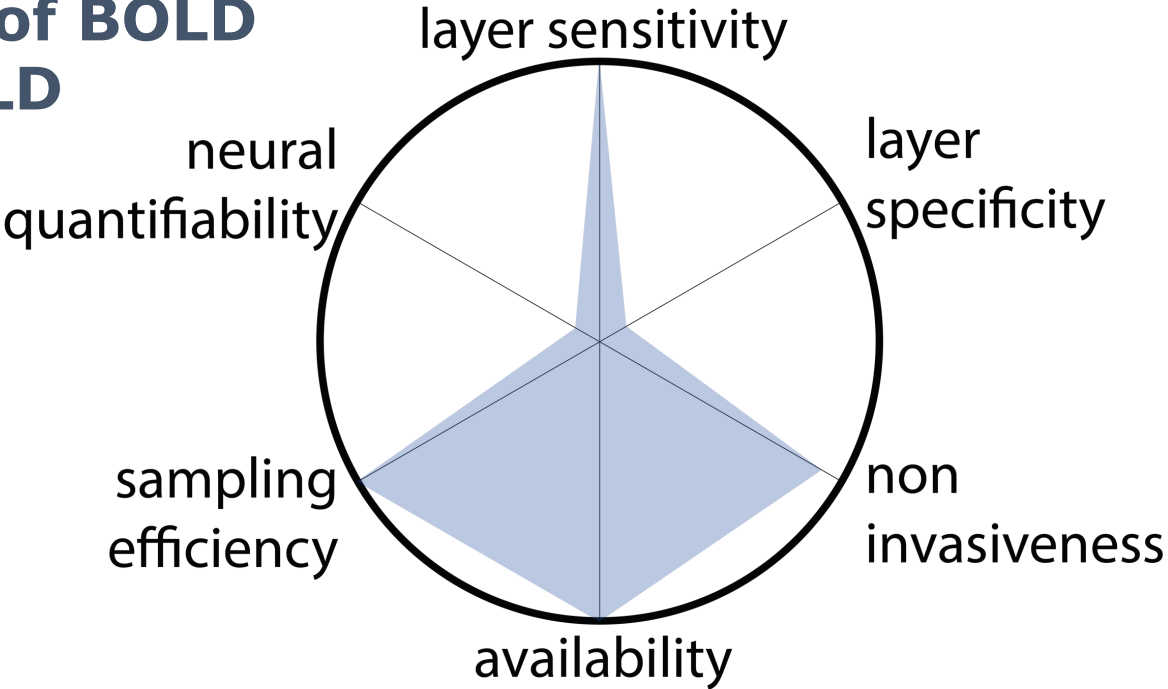
$$\frac{\Delta BOLD}{BOLD} = M \left[1 - \left(\frac{CMRO_2}{CMRO_2|_{rest}} \right)^\beta \left(\frac{CBV_v}{CBV_v|_{rest}} \right)^{\frac{\alpha_v - \beta}{\alpha_t}} \right]$$

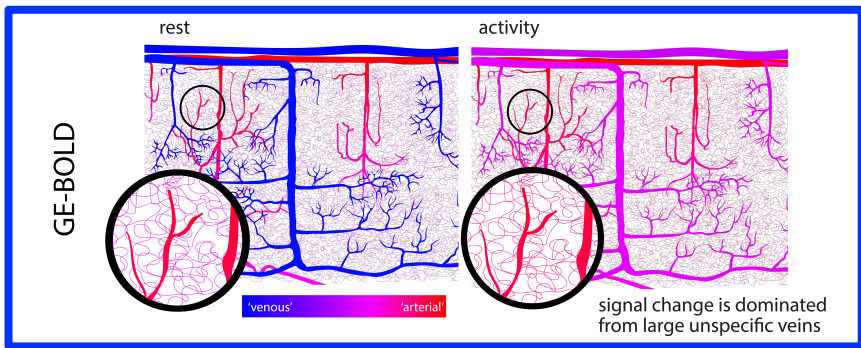


Sequence parameters:
 FOV: 4.8x4.8 cm²
 Resol [mm³]: **0.75x0.75x1.5**
 TR/TI1/TI2 = 3/0.95/2.45 s
TE = 18 ms
 7T

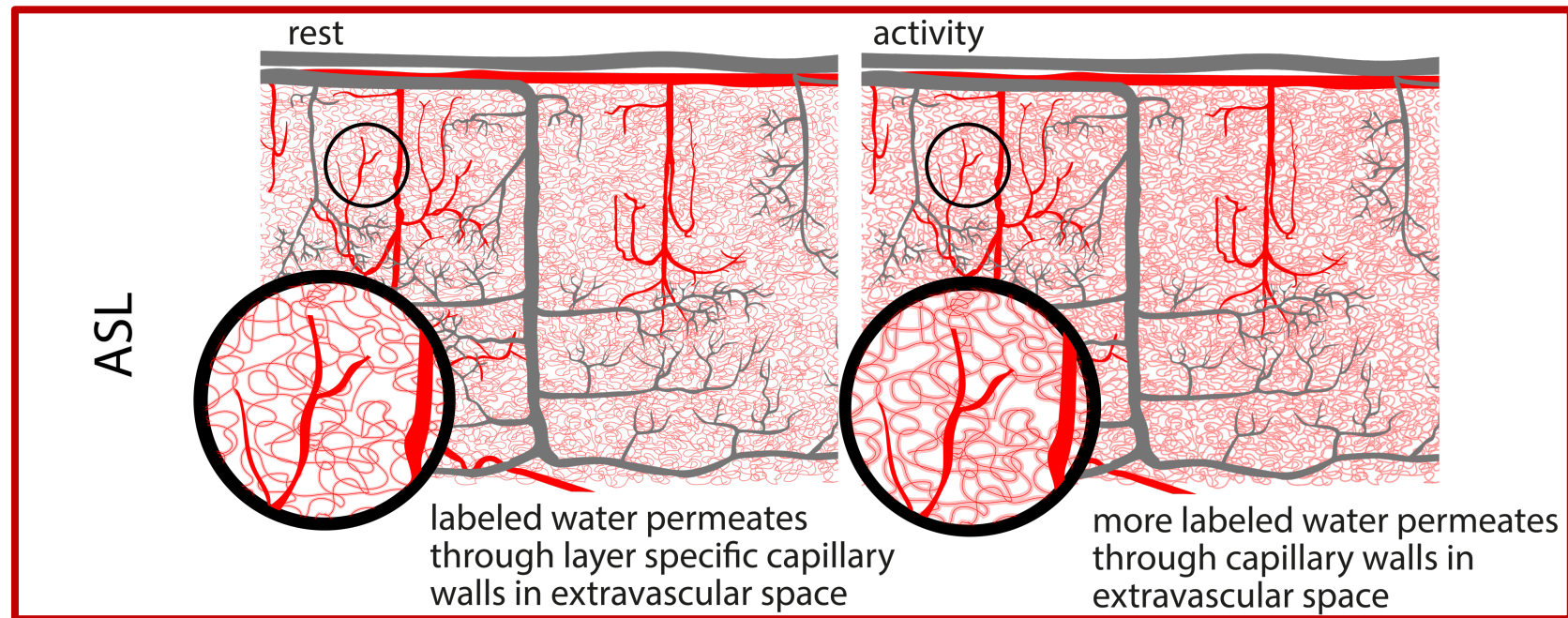
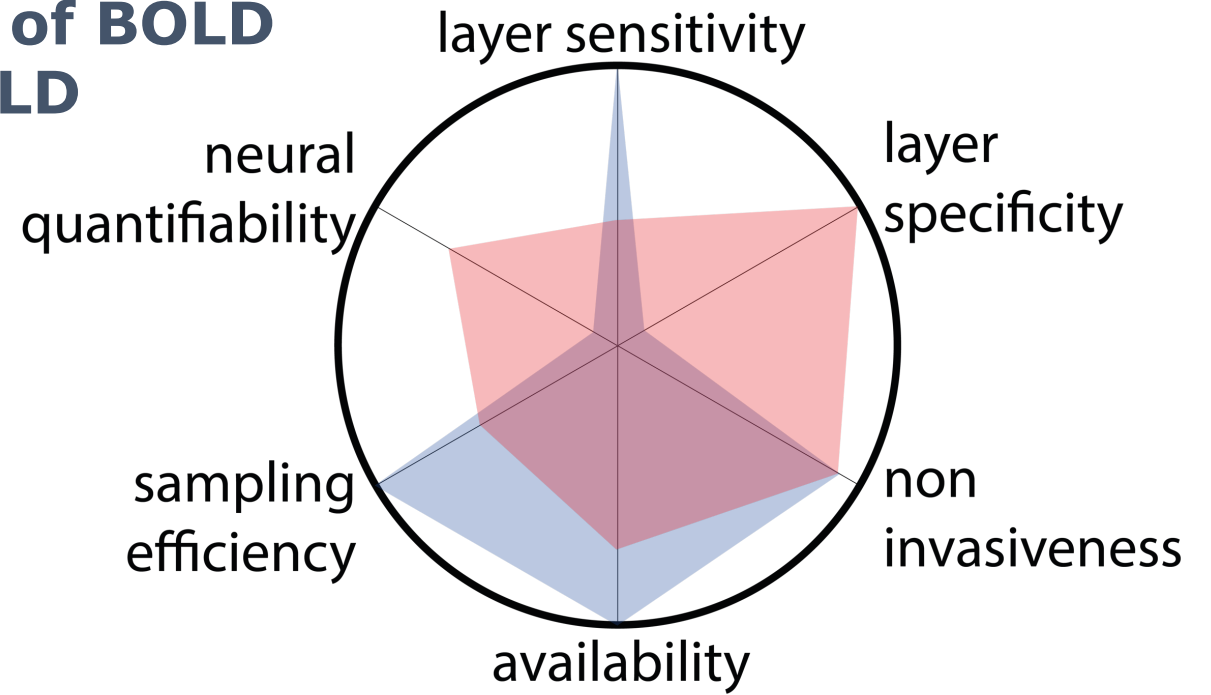


comparison of BOLD and non-BOLD methods

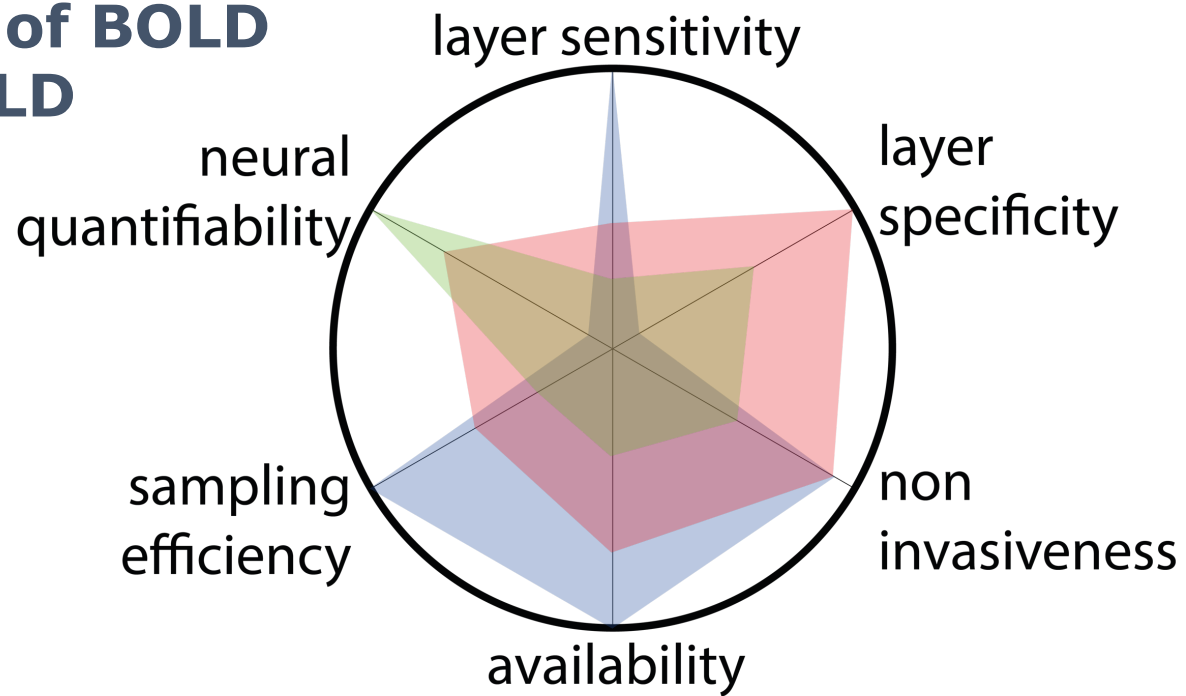
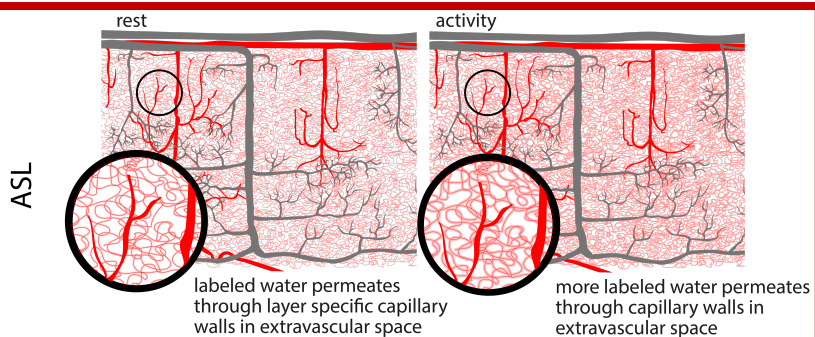
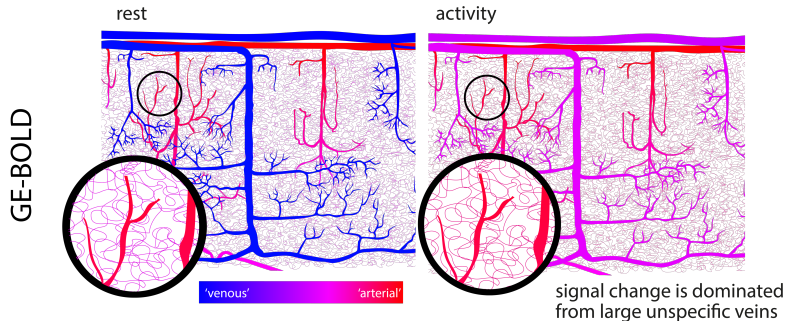




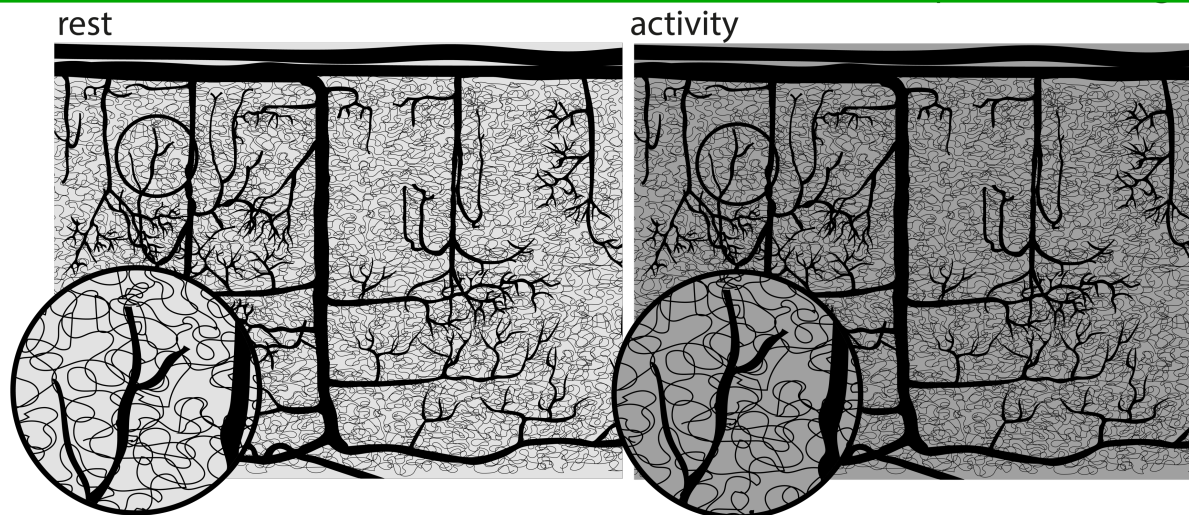
comparison of BOLD and non-BOLD methods



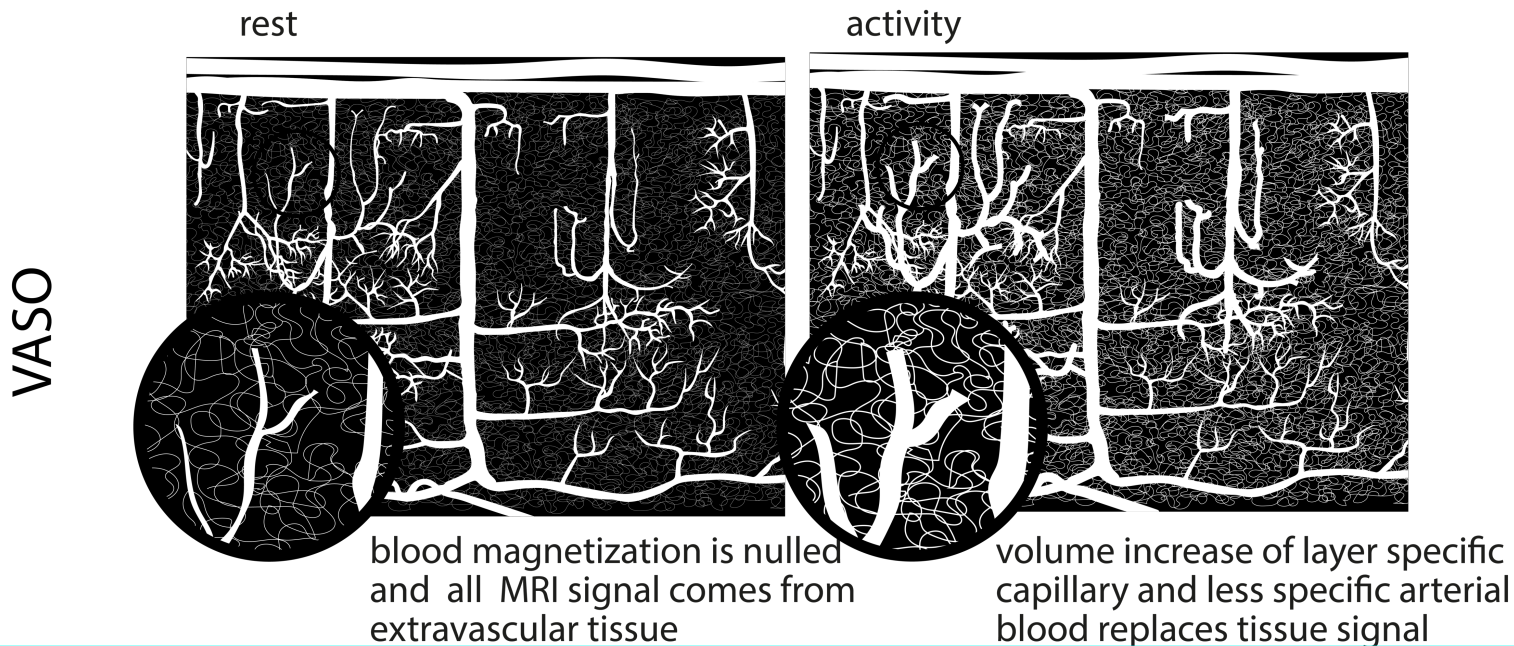
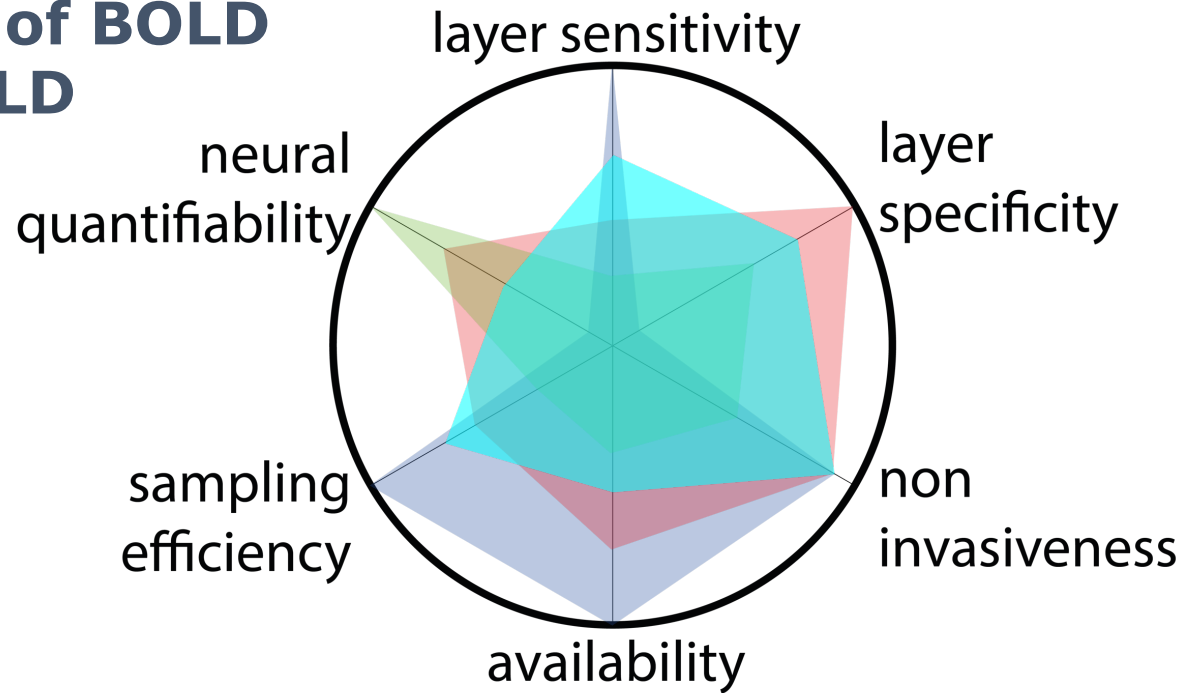
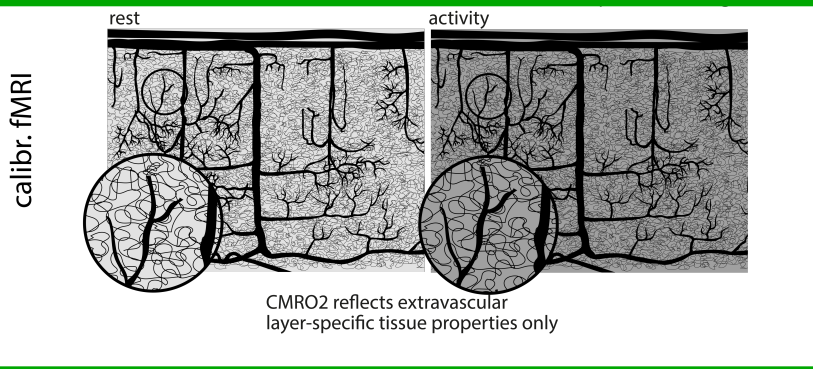
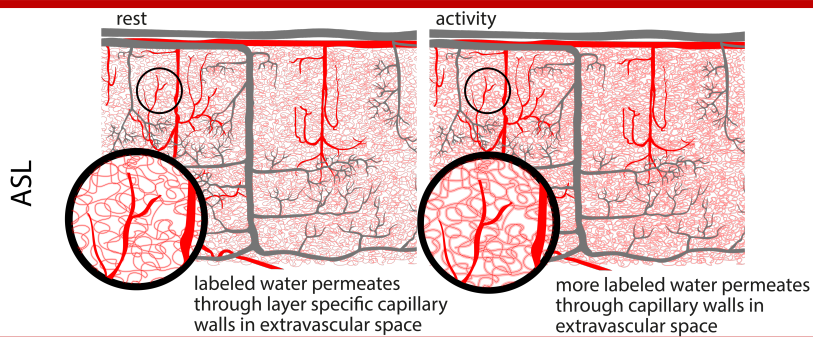
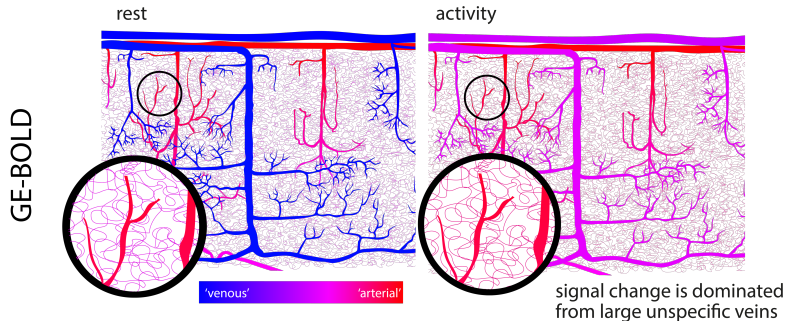
comparison of BOLD and non-BOLD methods



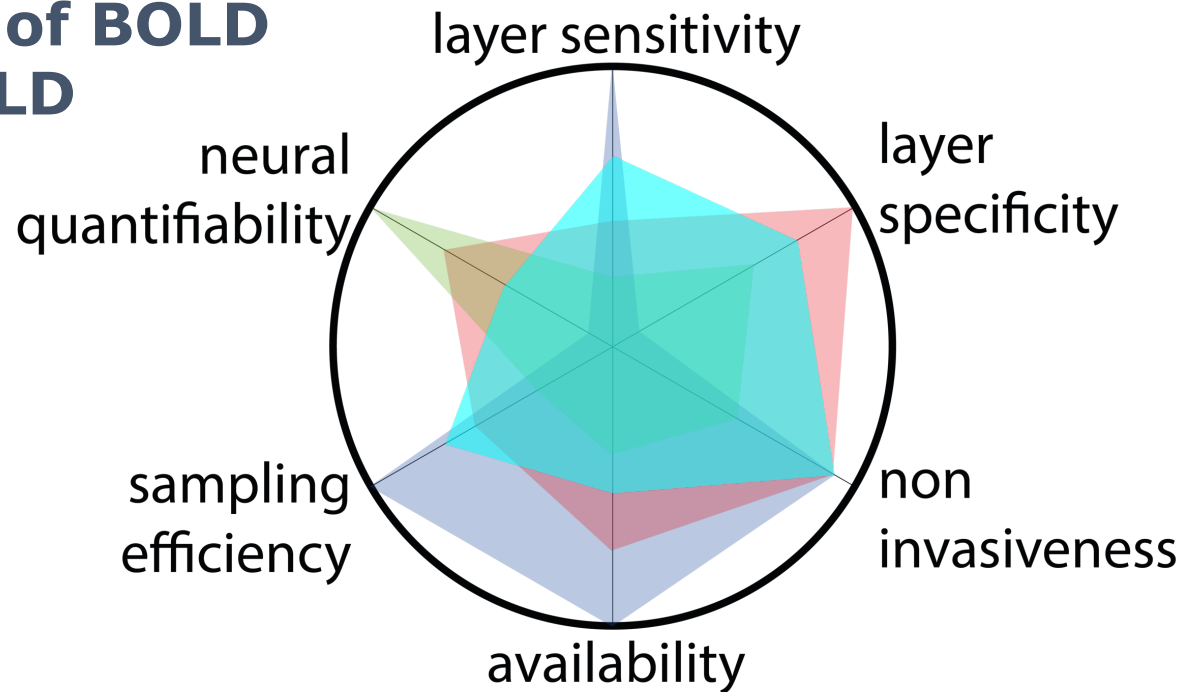
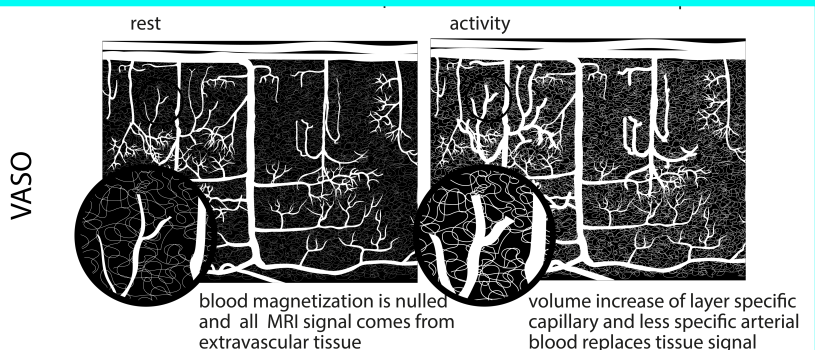
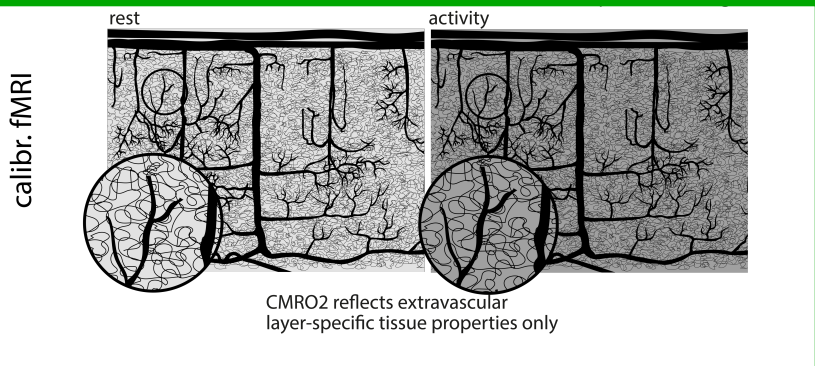
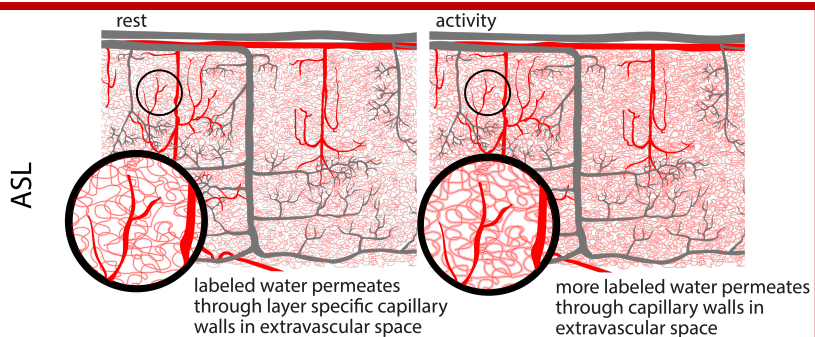
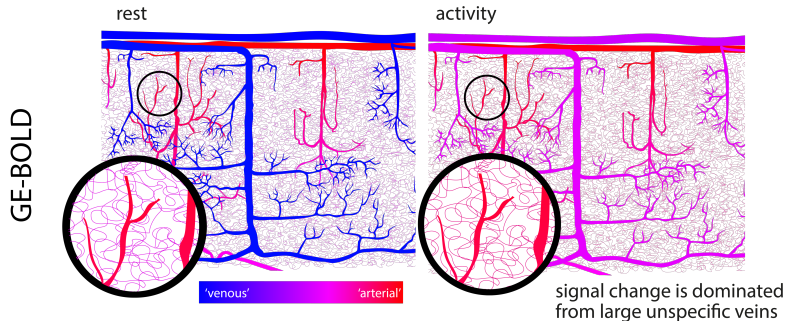
calibr. fMRI



comparison of BOLD and non-BOLD methods



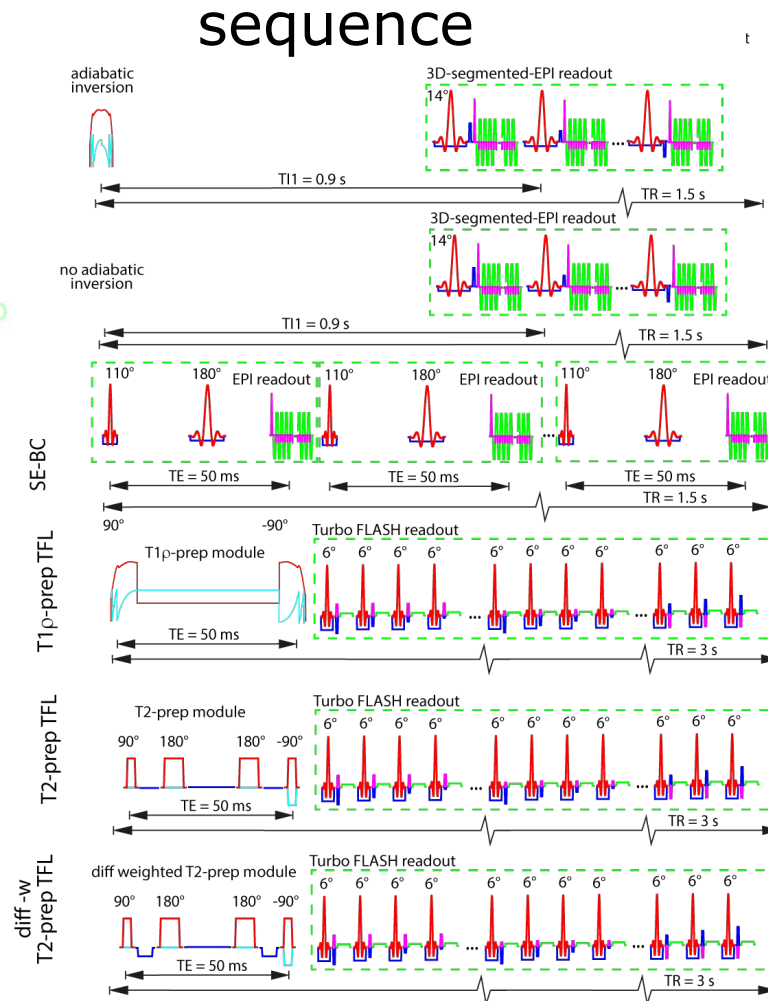
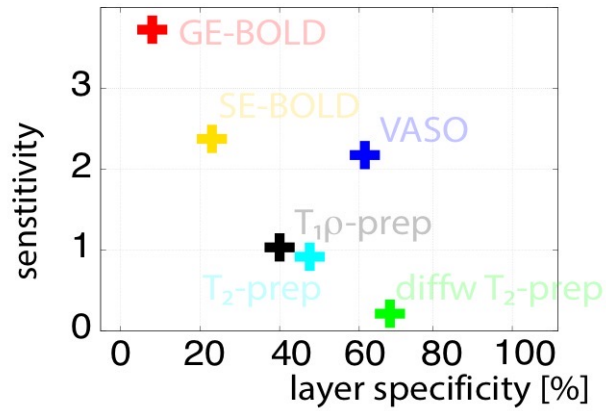
comparison of BOLD and non-BOLD methods



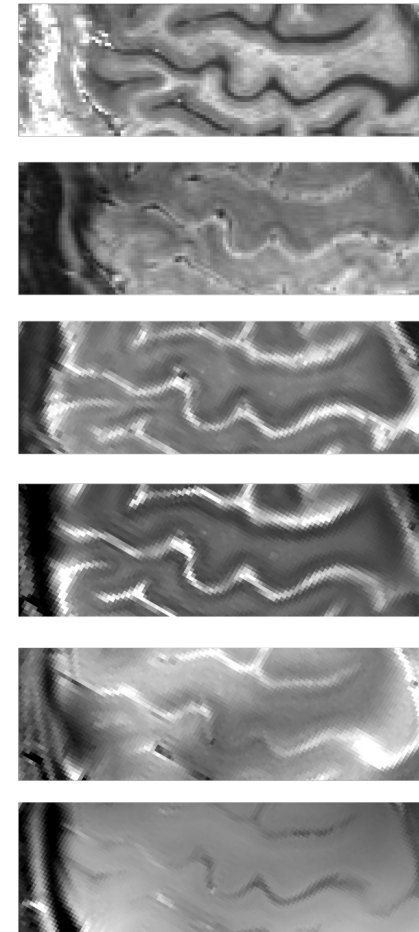
Summary:

- Non-BOLD layer-fMRI is noisier (reducing unwanted signal)
- Non-BOLD layer-fMRI is more specific
- Non-BOLD layer-fMRI is more quantitative

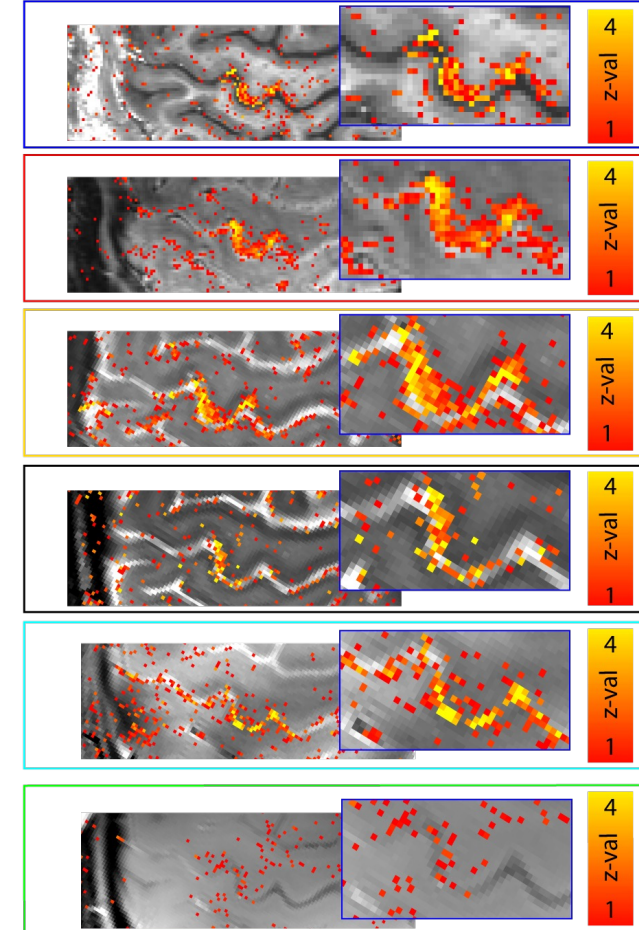
What about spin echo?



MRI contrast

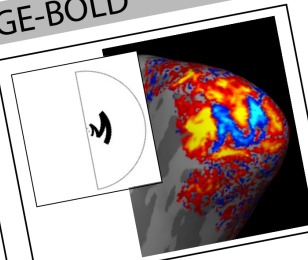


fMRI signal change



GE-BOLD vs. other sequences

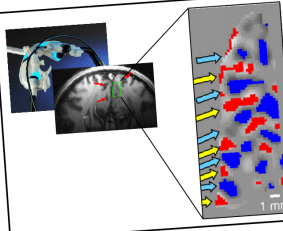
GE-BOLD



Sensitivity	★★★★★
Specificity	☆☆☆☆☆
Accessibility	★★★★★
Skill	☆☆☆☆☆
FOV and TR	★★★★★

Falmer 2010, NeuroImage

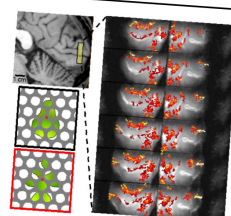
SE-BOLD



Sensitivity	★★★★★
Specificity	☆☆☆☆☆
Accessibility	★★★★★
Skill	☆☆☆☆☆
FOV and TR	★★★★★

Yacoub 2007, NeuroImage

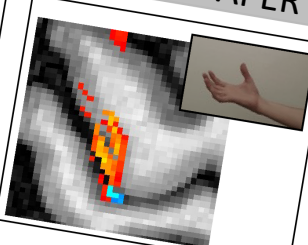
3D-GRASE



Sensitivity	★★★★★
Specificity	☆☆☆☆☆
Accessibility	☆☆☆☆☆
Skill	★★★★★
FOV and TR	☆☆☆☆☆

Zhang 2023


VASO & VAPER



Sensitivity	☆☆☆☆☆
Specificity	★★★★★
Accessibility	☆☆☆☆☆
Skill	☆☆☆☆☆
FOV and TR	★★★★★

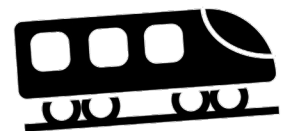
Huber 2017, Neuro

Airplane




Speed	★★★★★
Power	★★★★★
Comfort	☆☆☆☆☆
Size	☆☆☆☆☆
Price	★★★★★

Train




Speed	★★★★★
Power	☆☆☆☆☆
Comfort	☆☆☆☆☆
Size	★★★★★
Price	☆☆☆☆☆

Ship

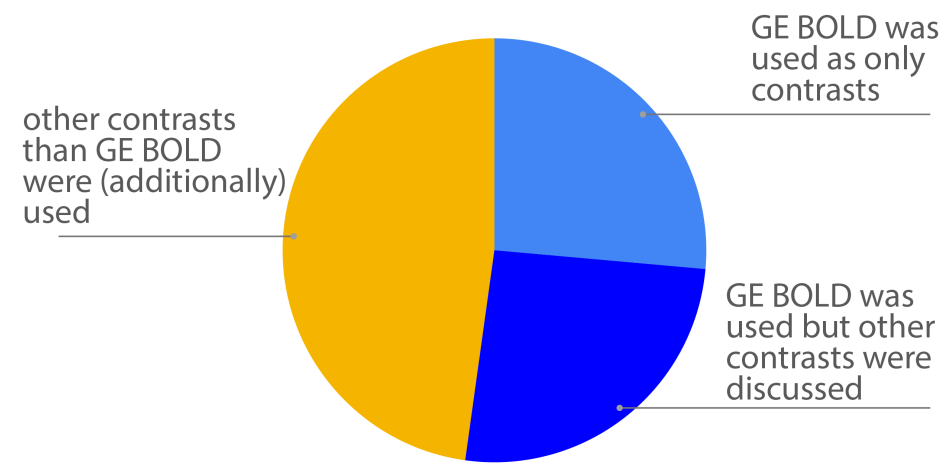


Speed	☆☆☆☆☆
Power	☆☆☆☆☆
Comfort	☆☆☆☆☆
Size	★★★★★
Price	☆☆☆☆☆

Car



Speed	☆☆☆☆☆
Power	★★★★★
Comfort	☆☆☆☆☆
Size	☆☆☆☆☆
Price	☆☆☆☆☆



layerfmri.com






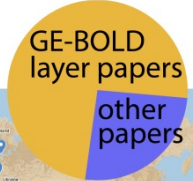
Oh these nouveau riche... So vulgar!

Spend all that money on Lamborghinis and then worry about food prices and end up at Mc-drive. Have some class and spend the money to eat something worth your money.

Feels like




7T Scanners



GE-BOLD layer papers

other papers



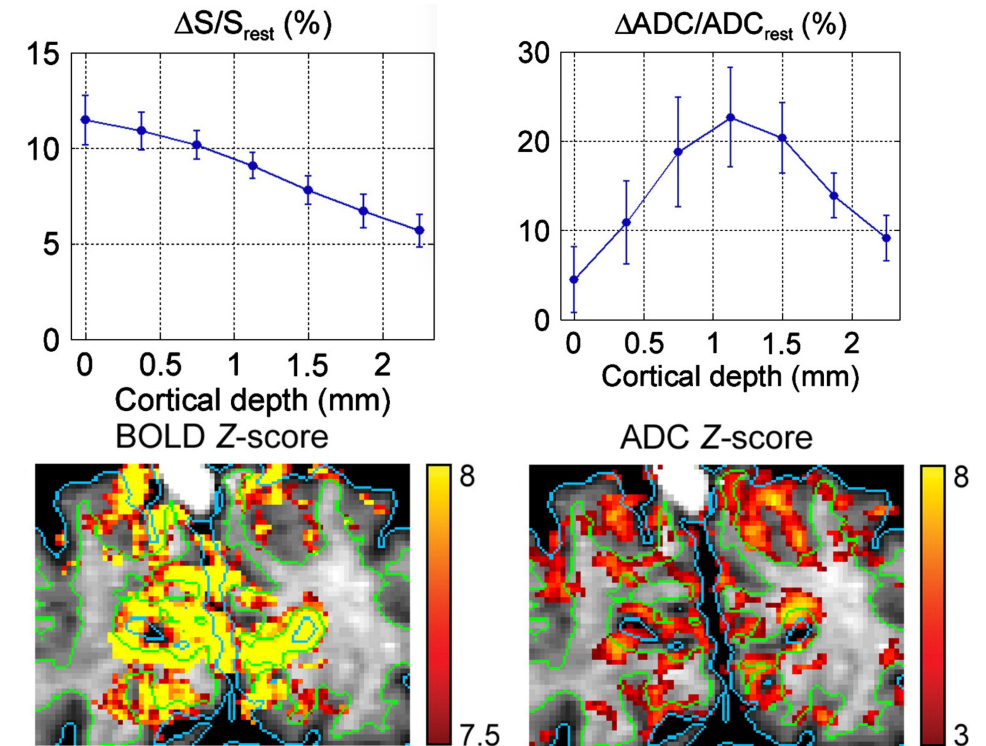
Oh these 'world leading' MRI centers... So cheap!

Spend all this money on 7Ts and then worry about sensitivity of fMRI contrasts and end up using GE-BOLD. Have some class and spend the money on long enough scan sessions to use VASO.

Other contrasts

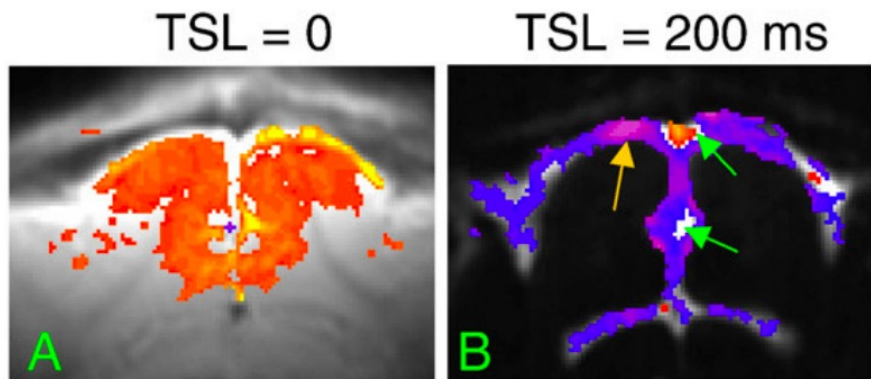


Allen Song et al. functional diffusion

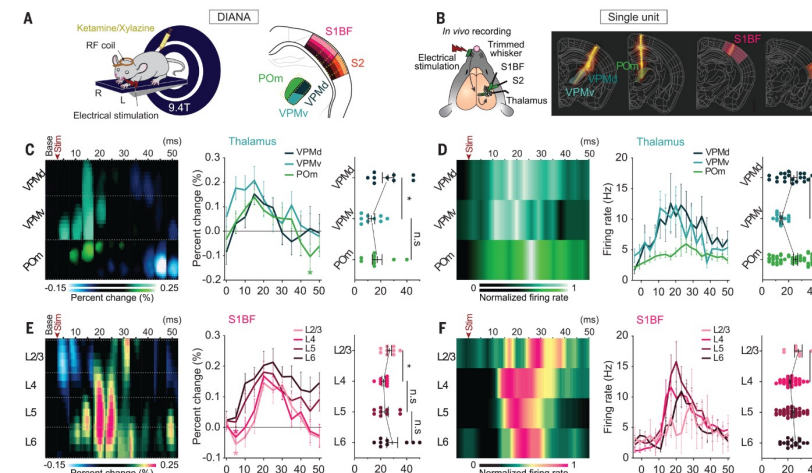


[Truong and Song 2009]

SG Kim et al.: functional $T_{1\rho}$ captures CSF volume change

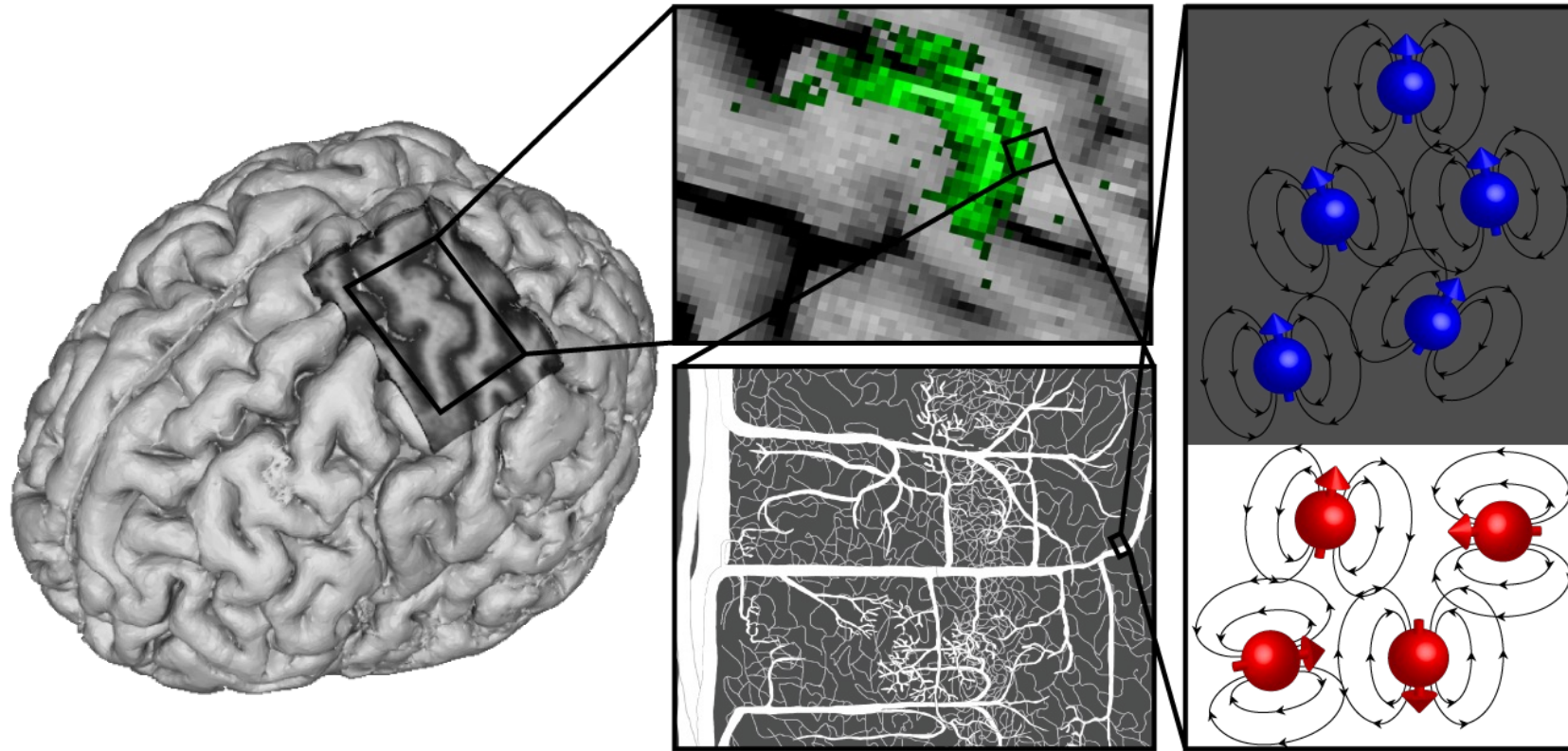


[Jin and Kim 2010]



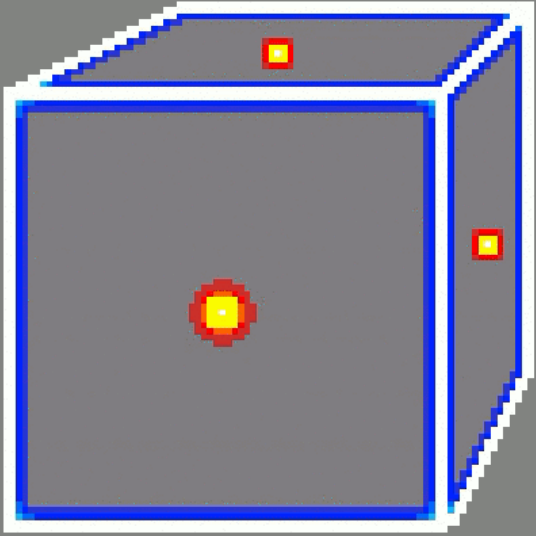
[Toi et al. 2022]

Functional contrast and processing strategies at high field and high resolution



Renzo (Laurentius) Huber

FMRI facility (FMRIF) at the NIMH/NINDS
of the National Institutes of Health (NIH)



LayNii:

a software suite for layer-fMRI

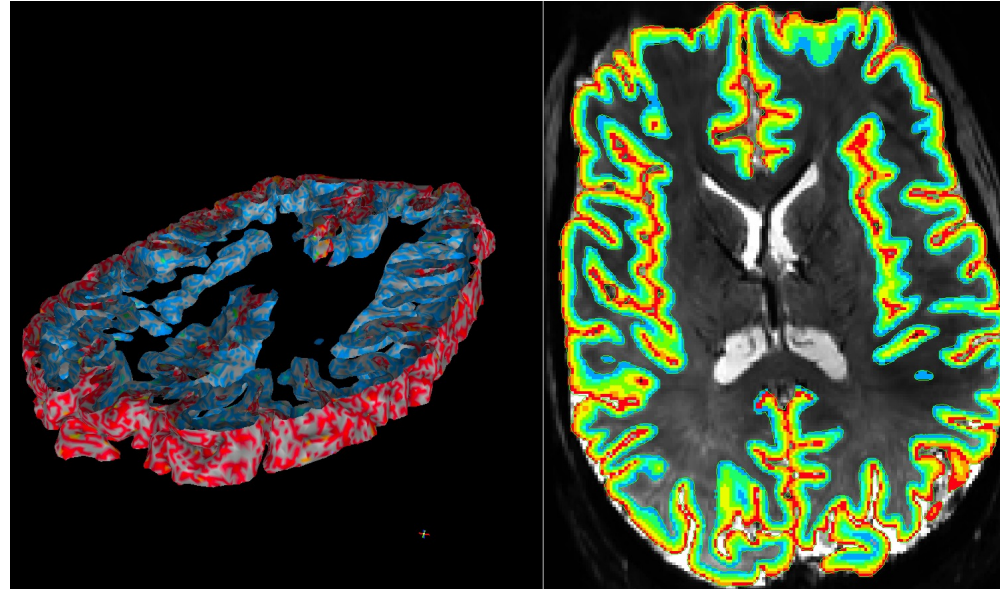
<https://github.com/layerfMRI/LAYNII>



Faruk Gulban

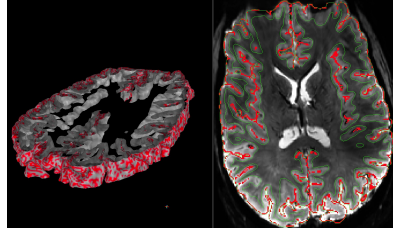
Why LAYNII

Small FOV complicates
surface-based
layerification



Why LAYNII

Small FOV complicates surface-based layerification

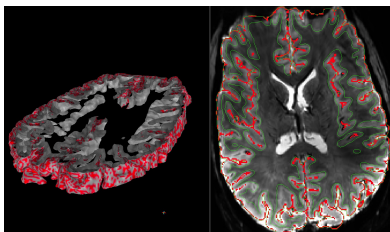


beer-wine dilemma



Why LAYNII

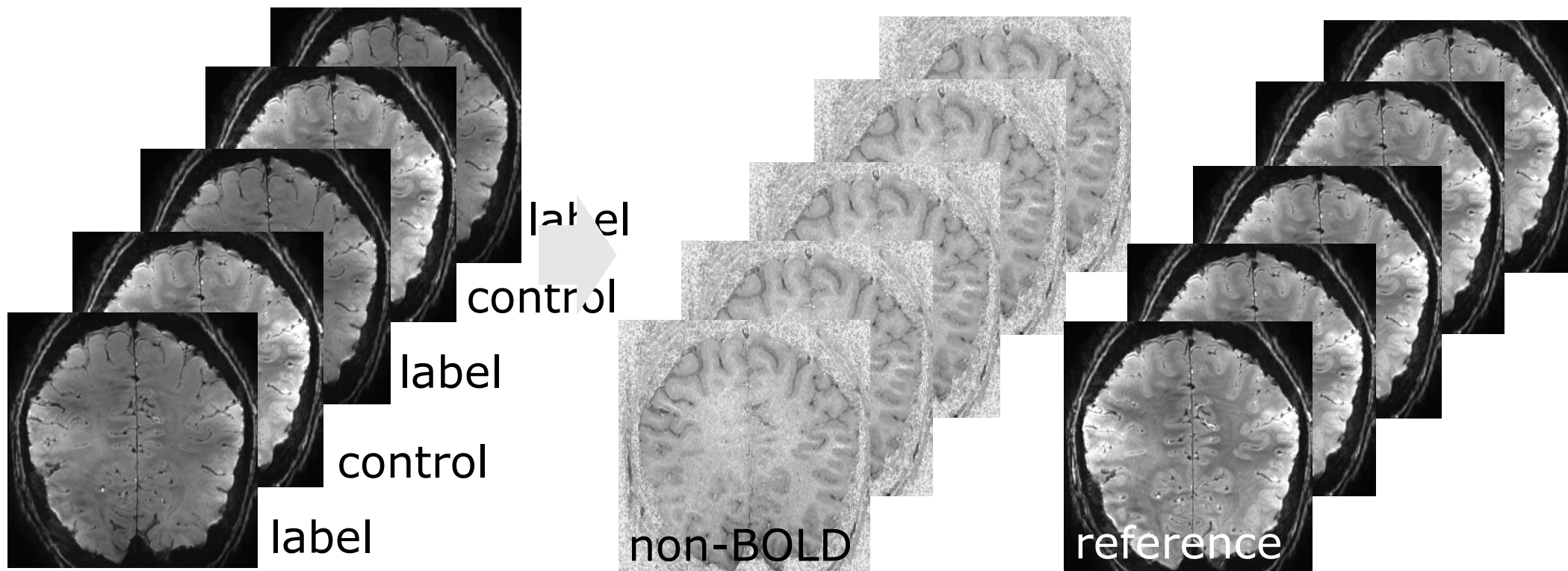
Small FOV complicates surface-based layerification



beer-wine dilemma

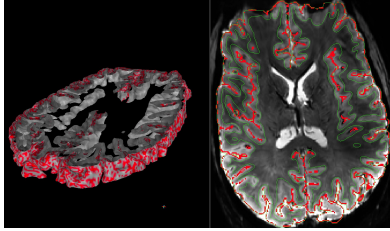


Non-BOLD contrasts



Why LAYNII

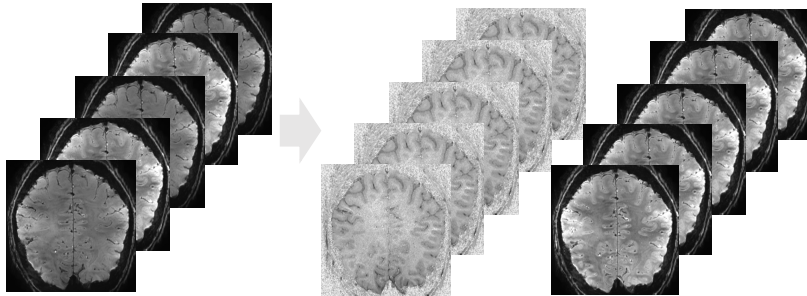
Small FOV complicates surface-based layerification



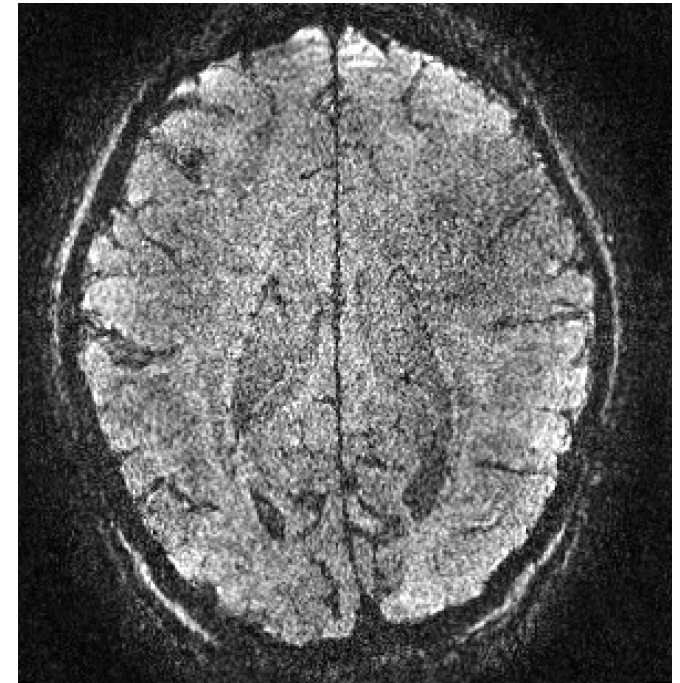
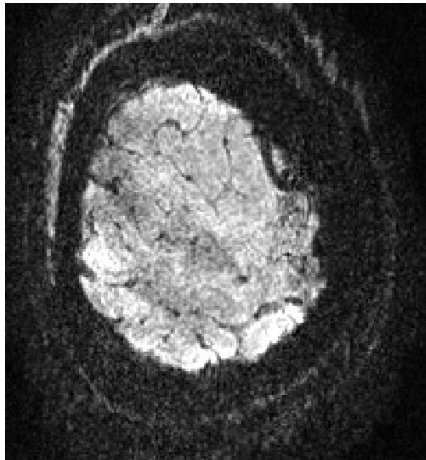
beer-wine dilemma



Non-BOLD contrasts

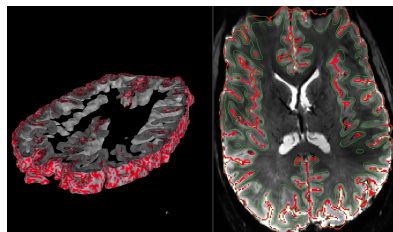


Artifacts, and noise



Why LAYNII

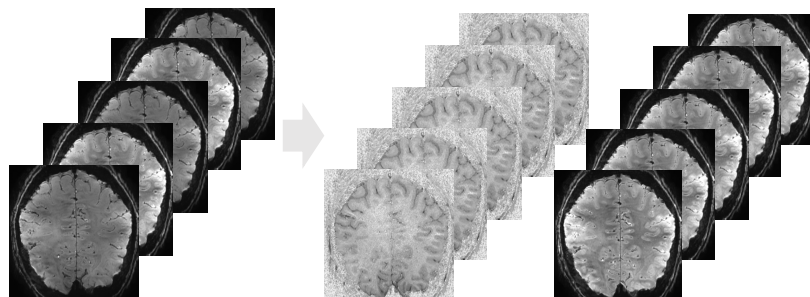
Small FOV complicates surface-based layerification



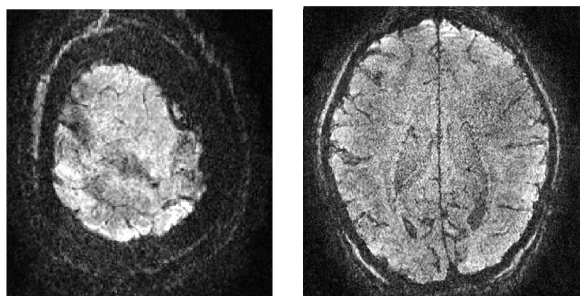
beer-wine dilemma



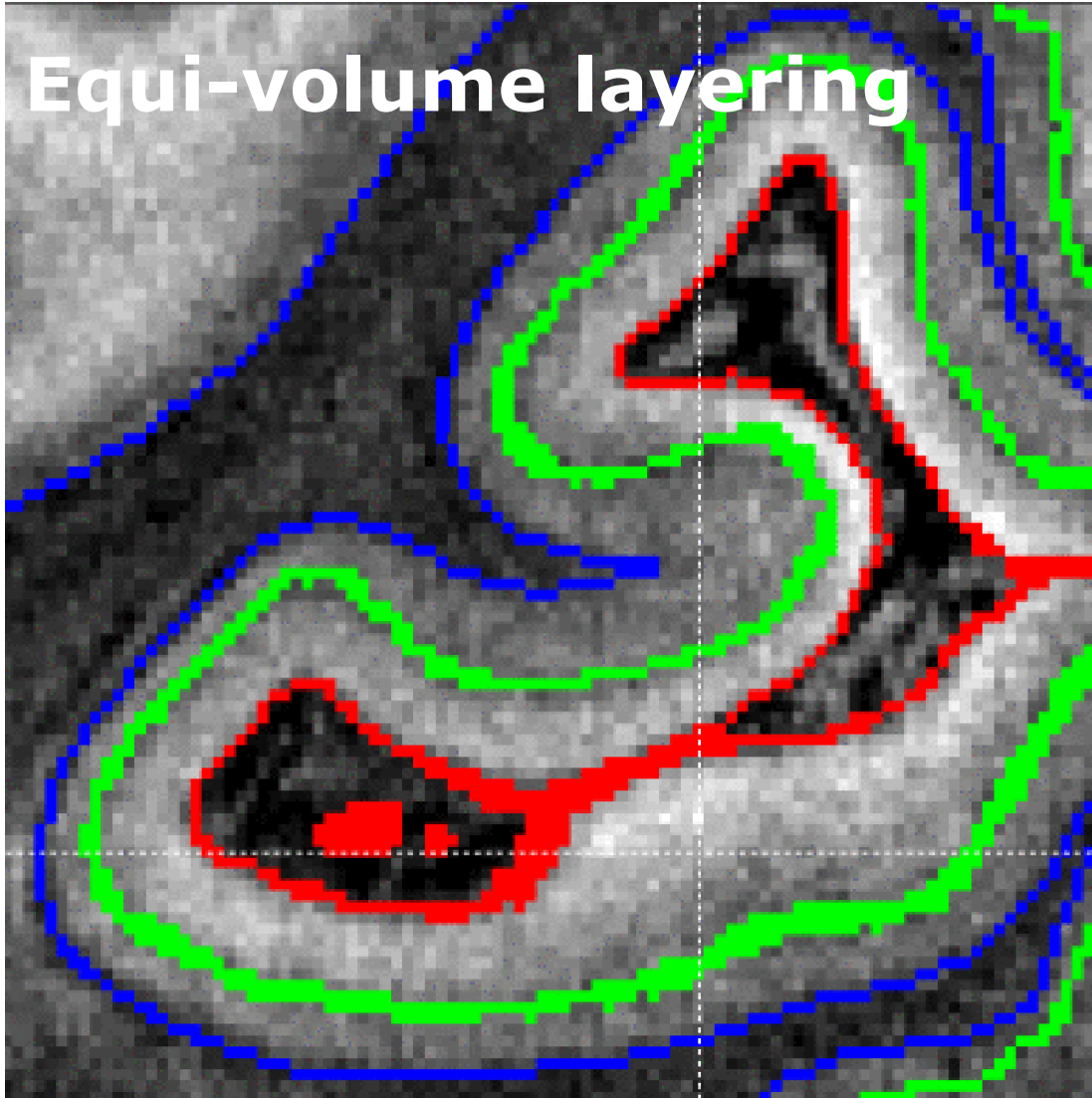
Non-BOLD contrasts



Artifacts, and noise



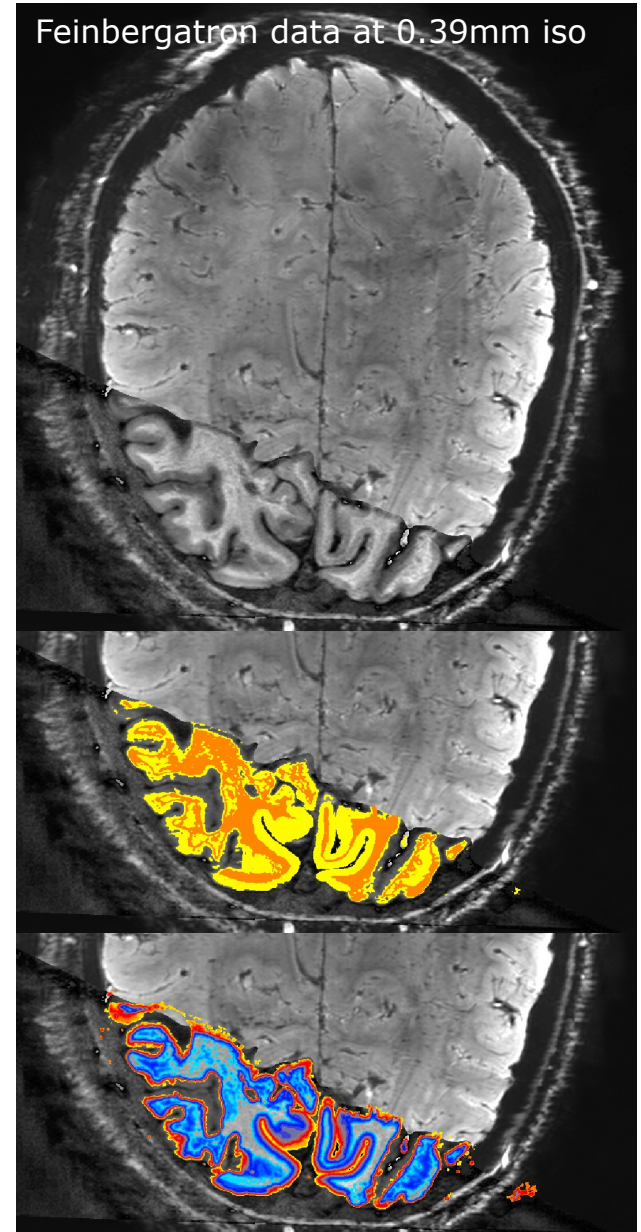
Layerification with the LayNii program LN2_LAYERS



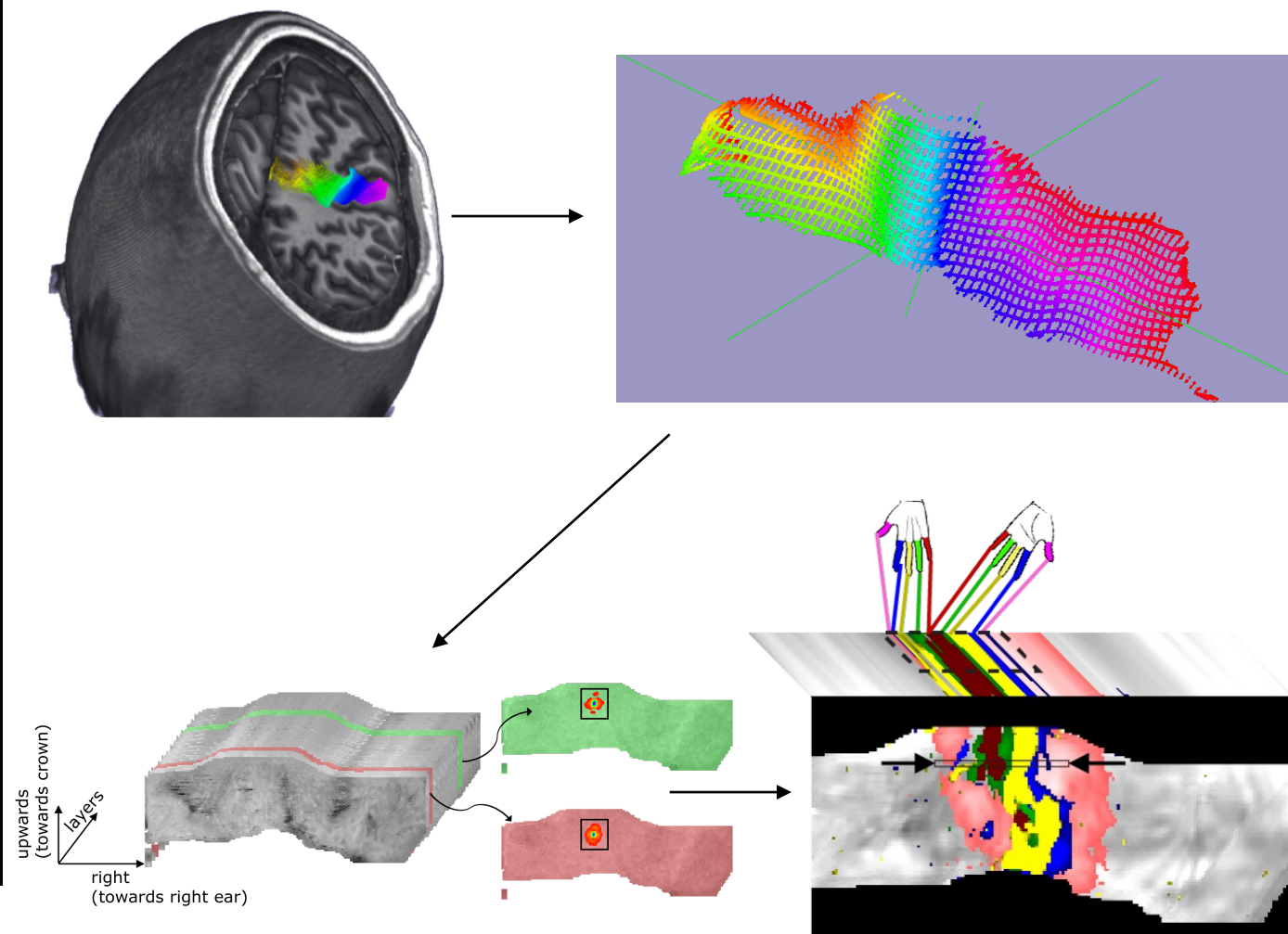
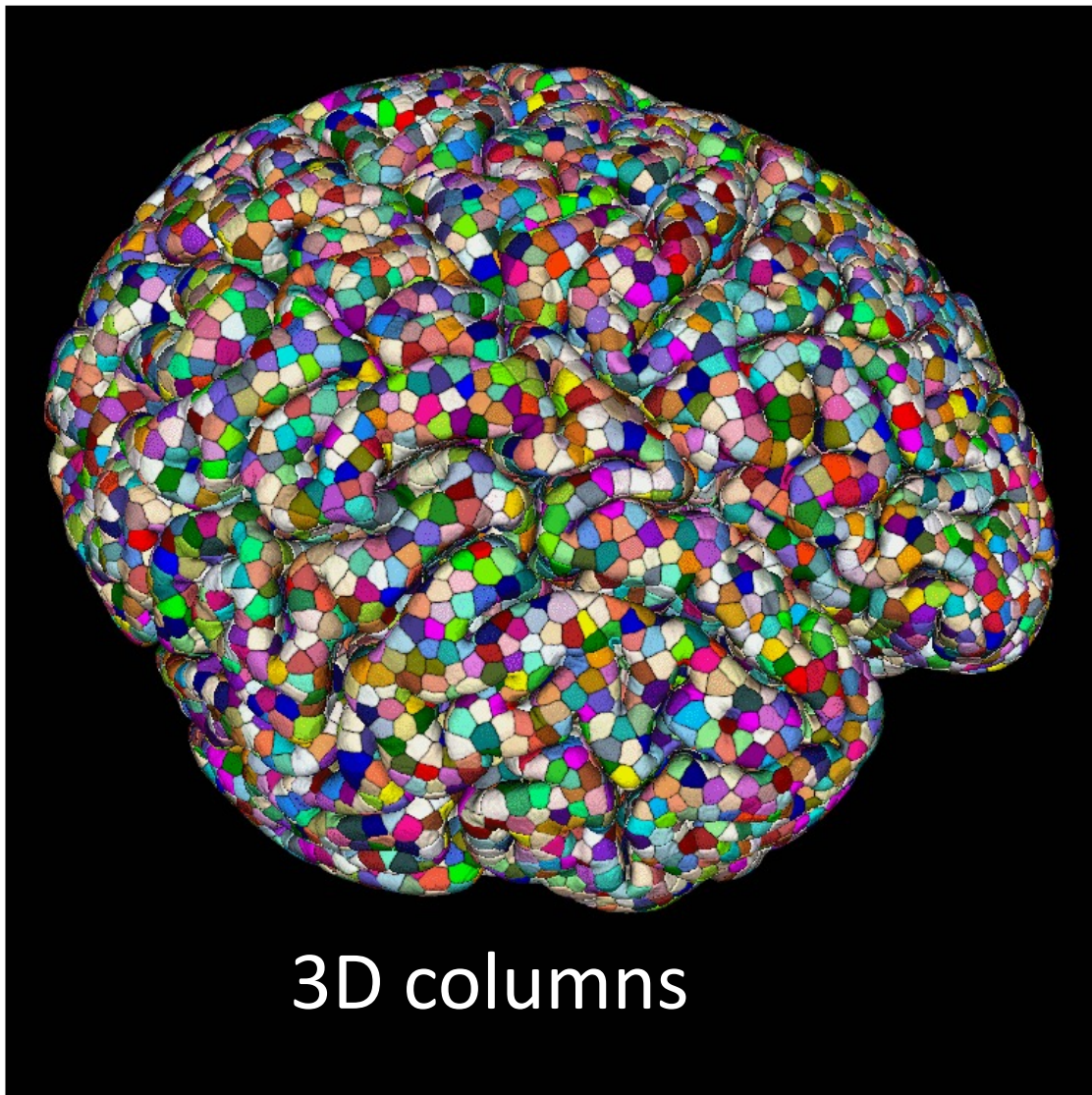
Ding et al. (2016), T2*-weighted, small section of V1.

See also:

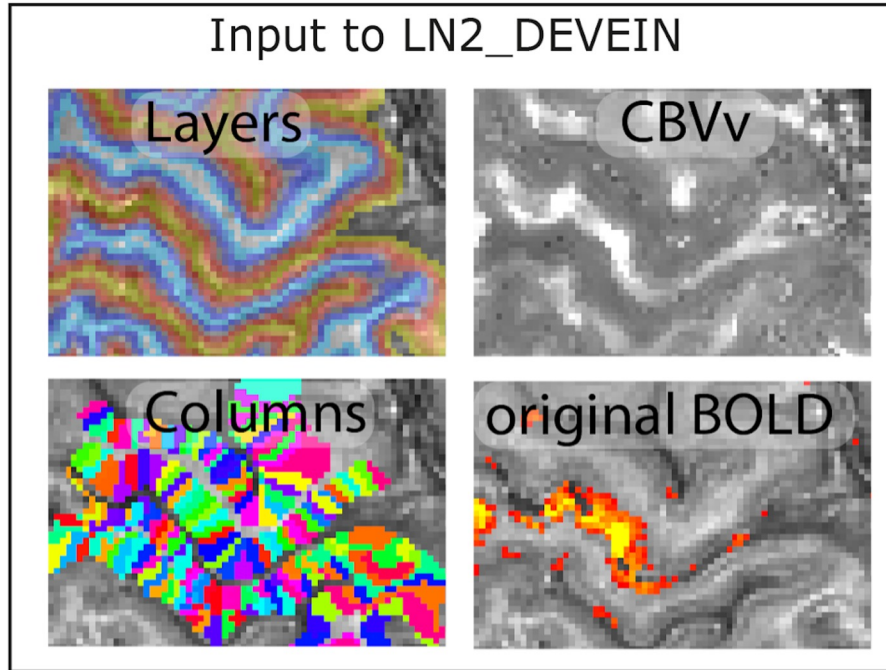
- Waehnert et al. 2014
- Kemper et al., 2018
- Wagstyl et al., 2020
- Bazin et al., 2014
- Huntenburg et al., 2018
- Laminar Python 2017
- Van Mourik et al., 2019
- Wagstyl et al. (2020),



Column generation



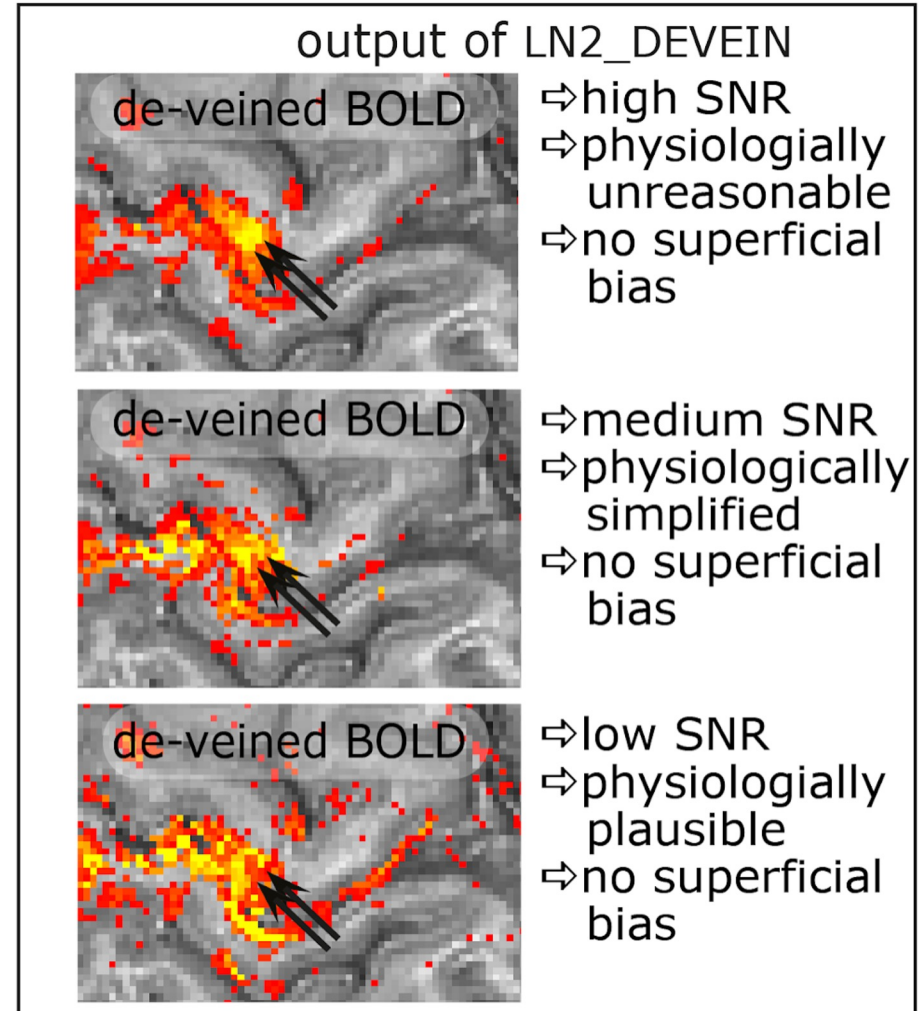
De-veining (a.k.a. model based GE-BOLD vein-effect mitigation)



linear offset correction

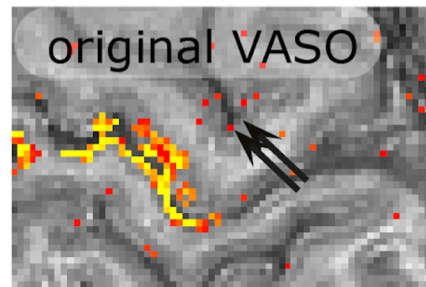
CBVv gain correction

leakage deconvolution



Heinze et al. 2016,
Markuerkiaga et al. 2016,
Marquardt et al., 2018,
Merola et al. 2018,
Corbitt et al. 2018,
Havlicek et al. 2019,
Puckett et al. 2016.

for reference



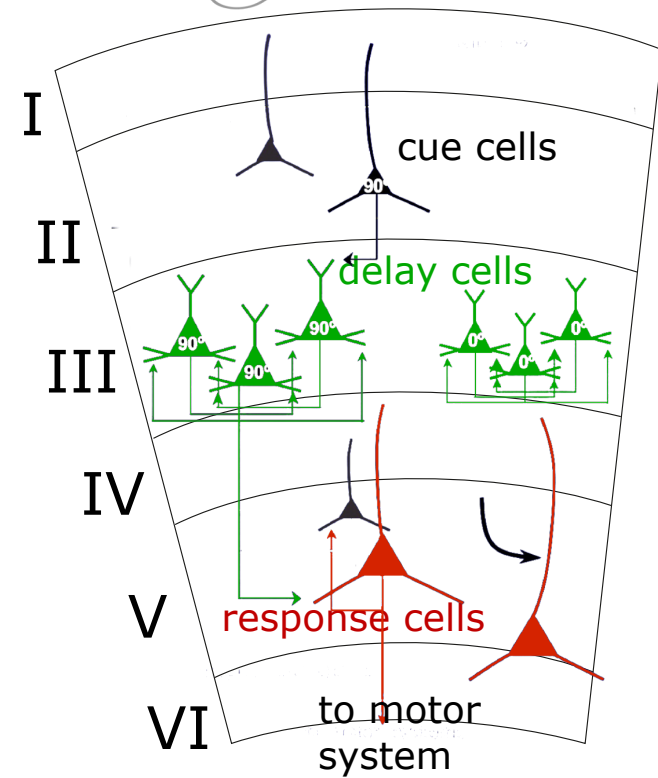
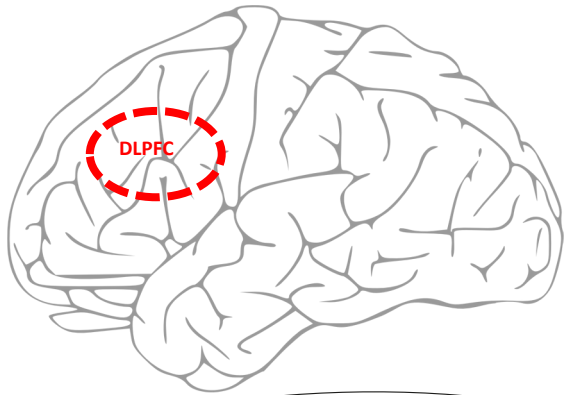
Layer-specific smoothing



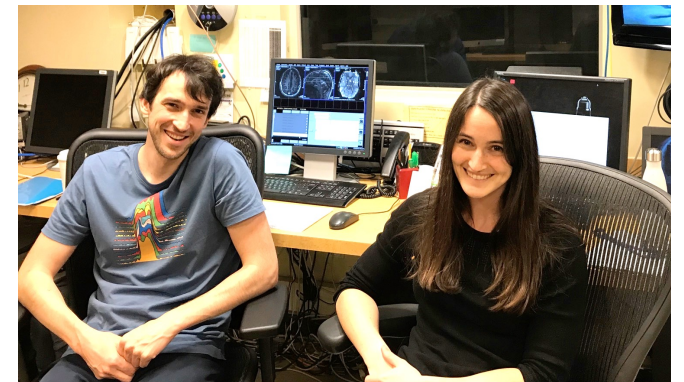
See also Blazejewska et al., 2019

layer-fMRI (post) processing

Most common layer-fMRI: locating activity changes by subtracting task conditions

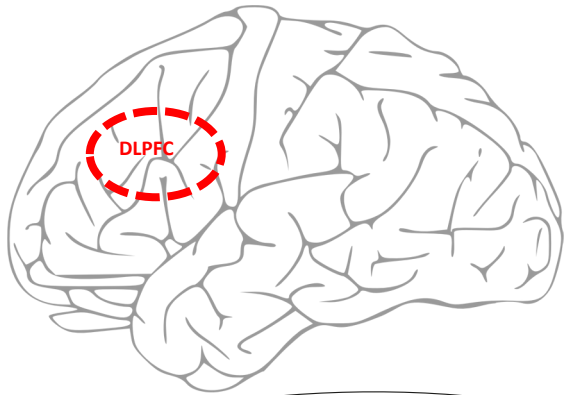


*Adapted from Arnsten et al., 2012
Wang et al., 2004; Opris et al., 2011;
Markowitz et al., 2015; Bastos et al., 2018*



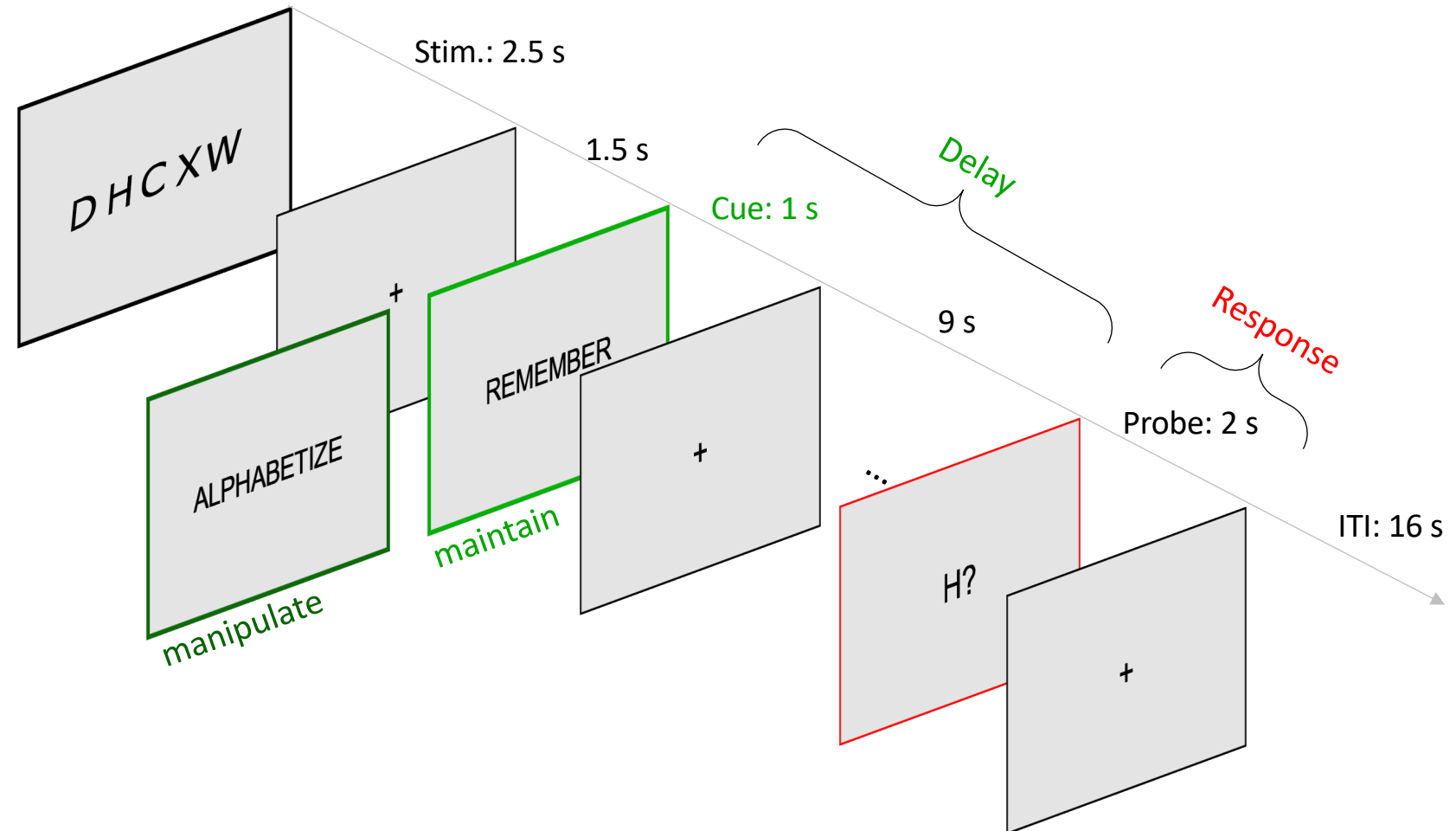
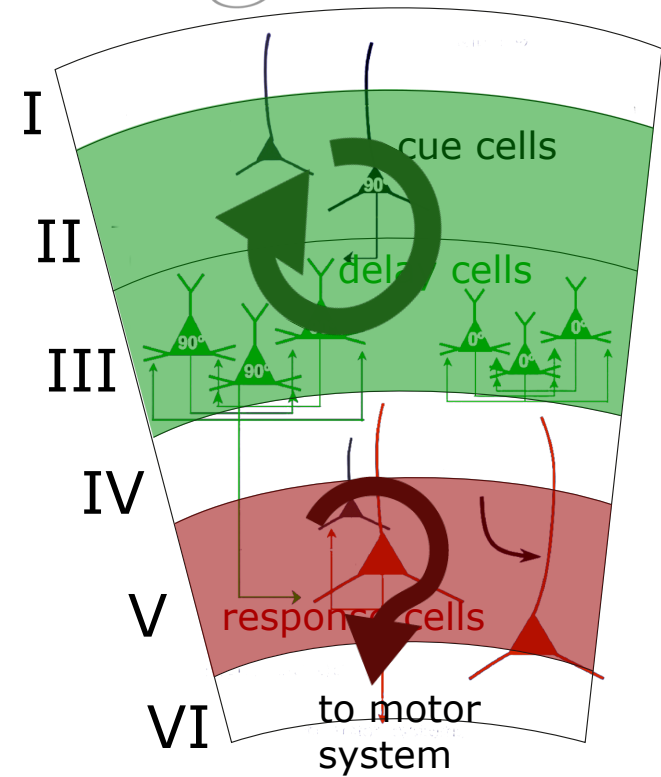
Emily Finn [2019]

Most common layer-fMRI: locating activity changes by subtracting task conditions



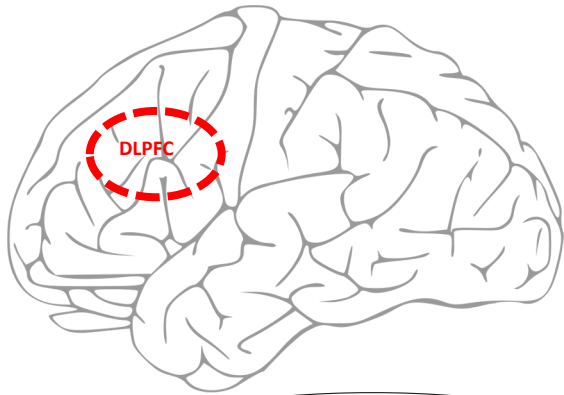
Task: manipulation vs. maintenance
response vs. letter sorting

[D'Esposito et al., 1999]

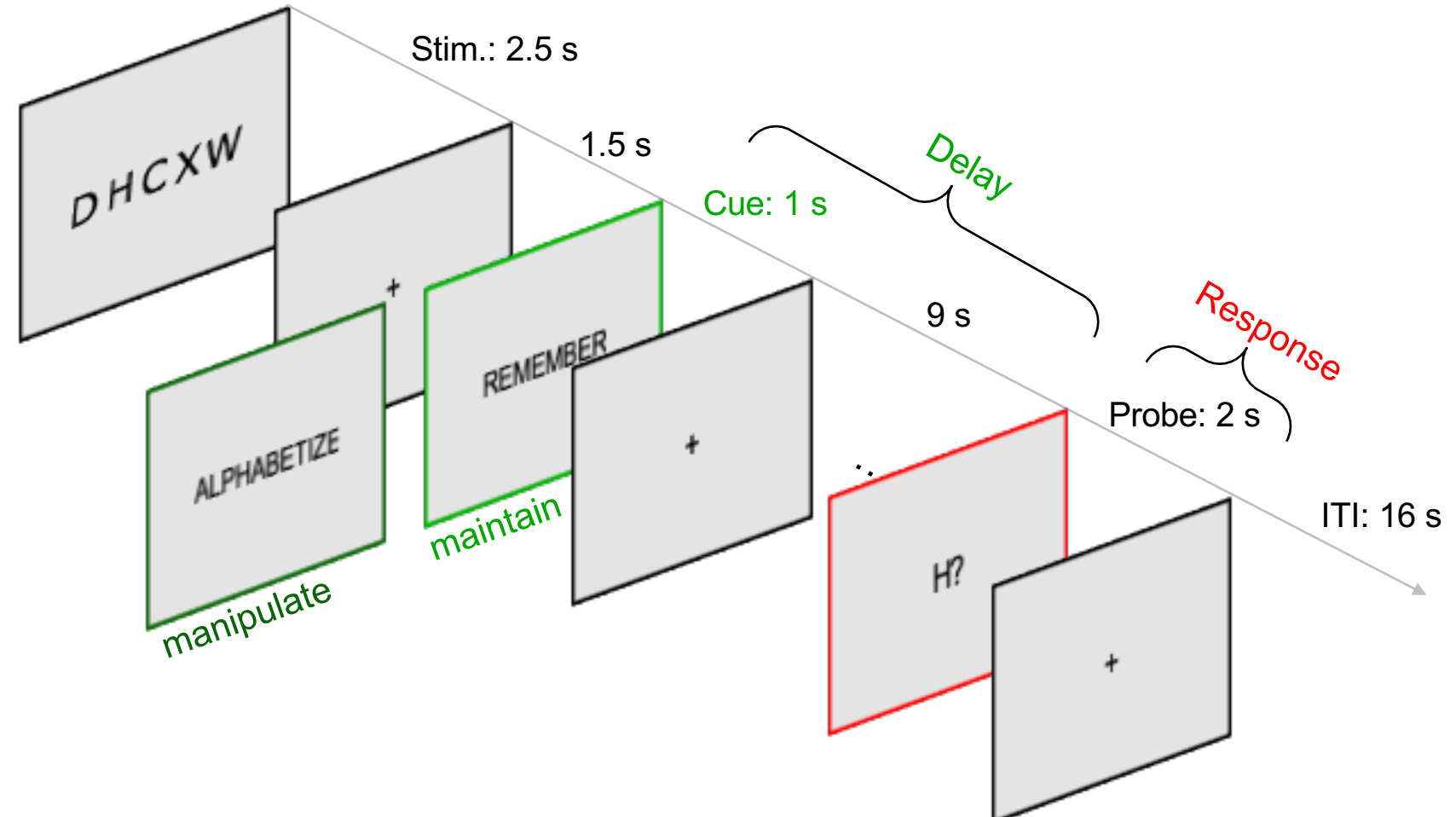
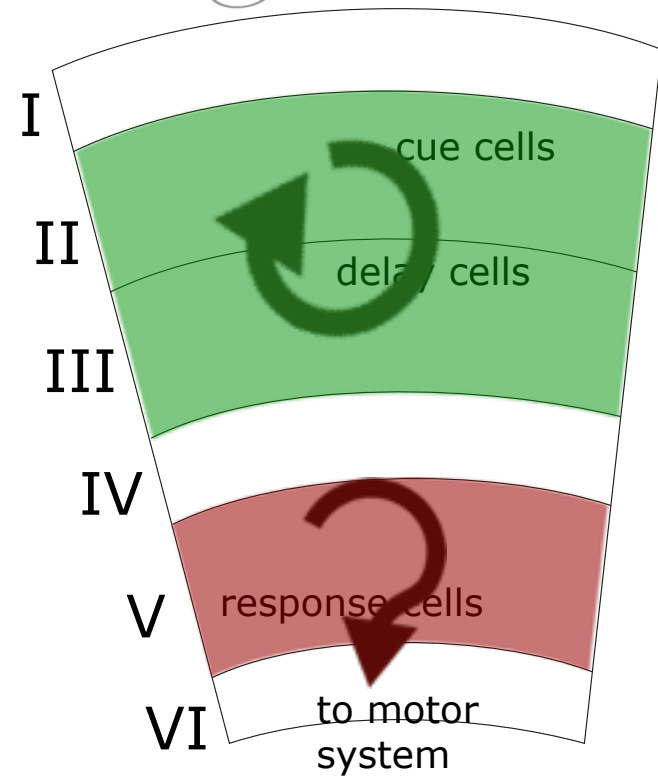


Adapted from Arnsten et al., 2012
Wang et al., 2004; Opris et al., 2011;
Markowitz et al., 2015; Bastos et al., 2018

Most common layer-fMRI: locating activity changes by subtracting task conditions

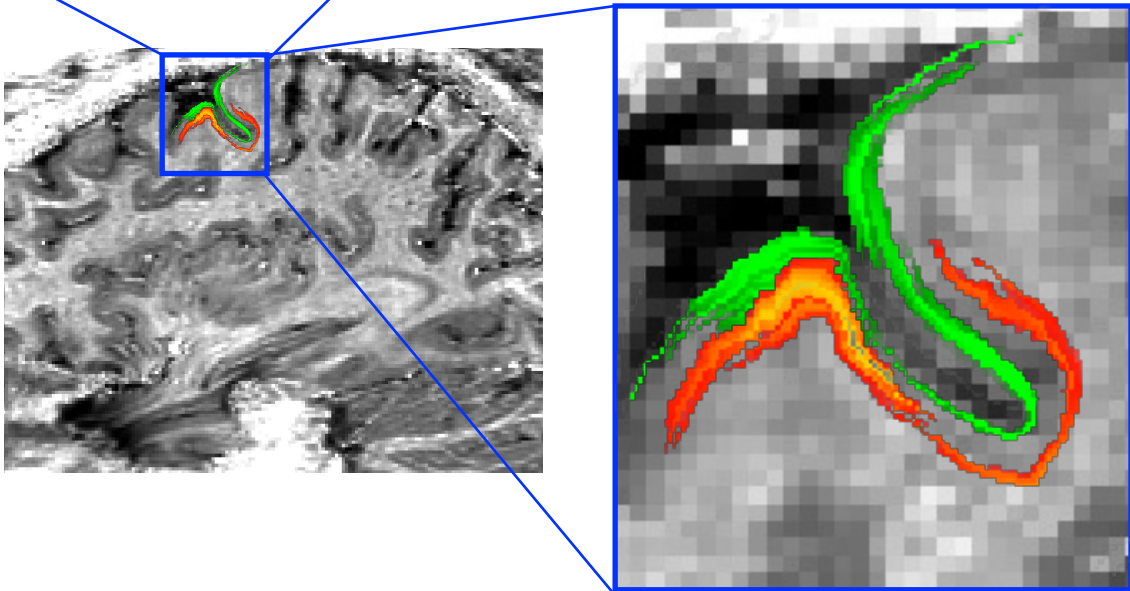
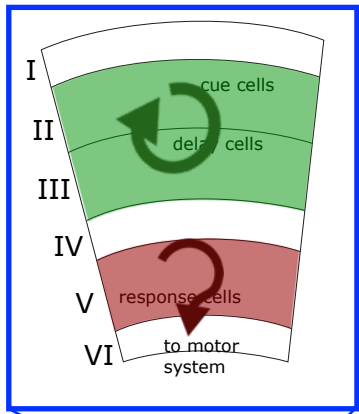


Task: manipulation vs. maintenance
response vs. letter sorting

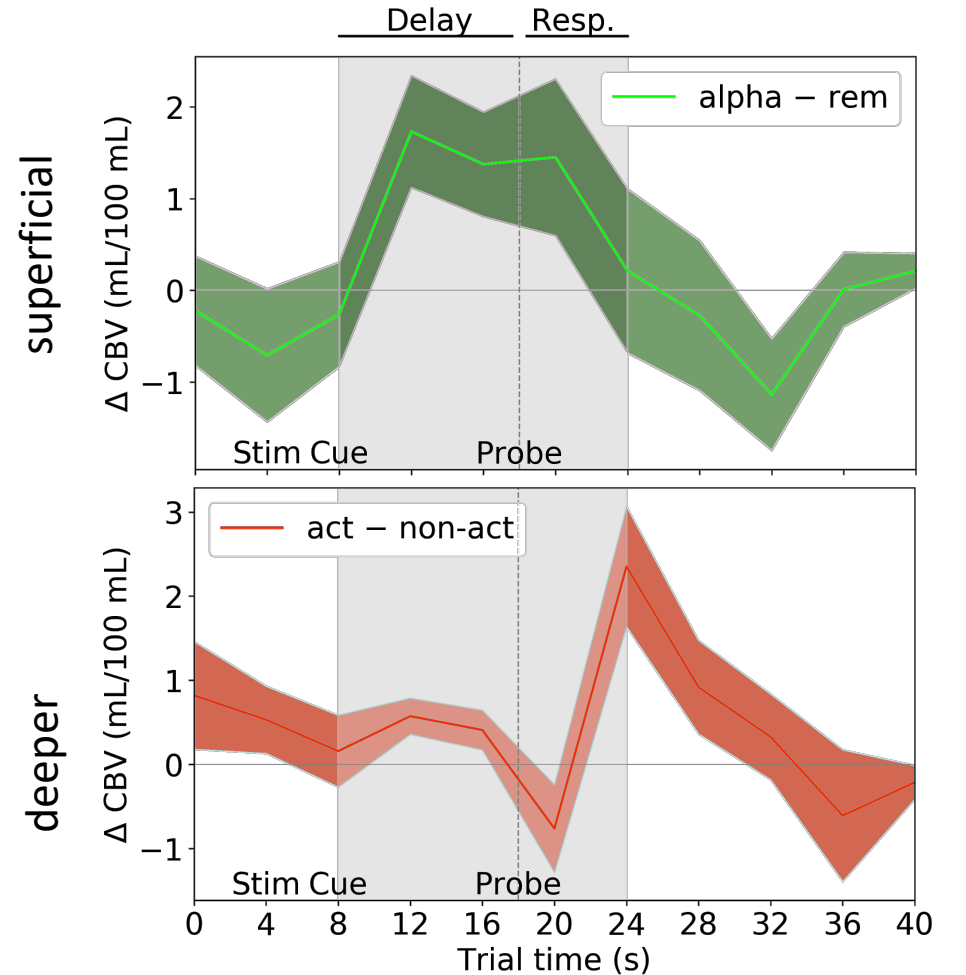
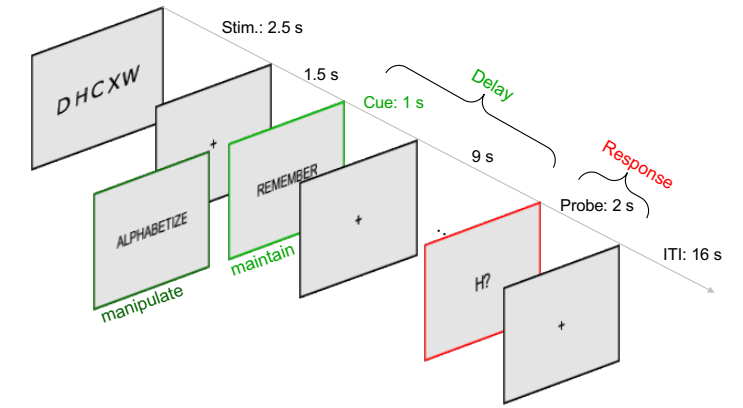


Adapted from Arnsten et al., 2012
Wang et al., 2004; Opris et al., 2011;
Markowitz et al., 2015; Bastos et al., 2018

Results

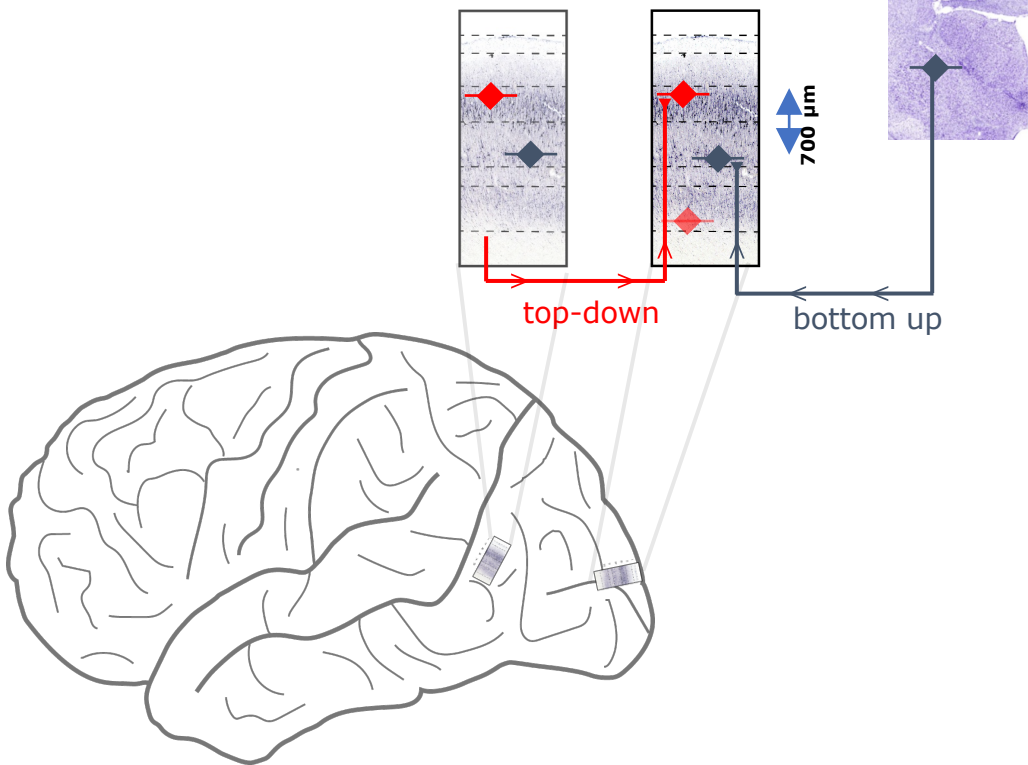


responding vs. rest		
0.2	Δ CBV [ml]	1.5
alphabetizing vs. remembering		
0.2	Δ CBV [ml]	0.8

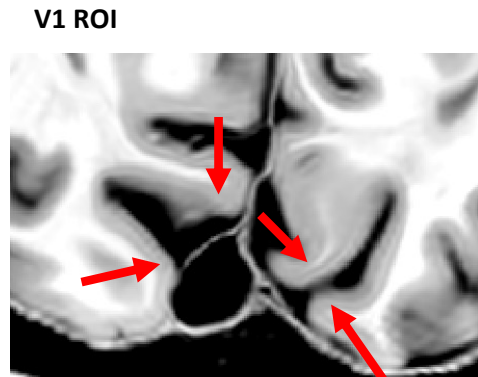
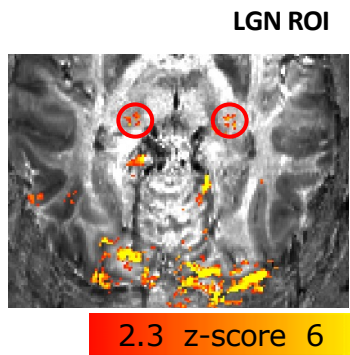


higher hierarchy

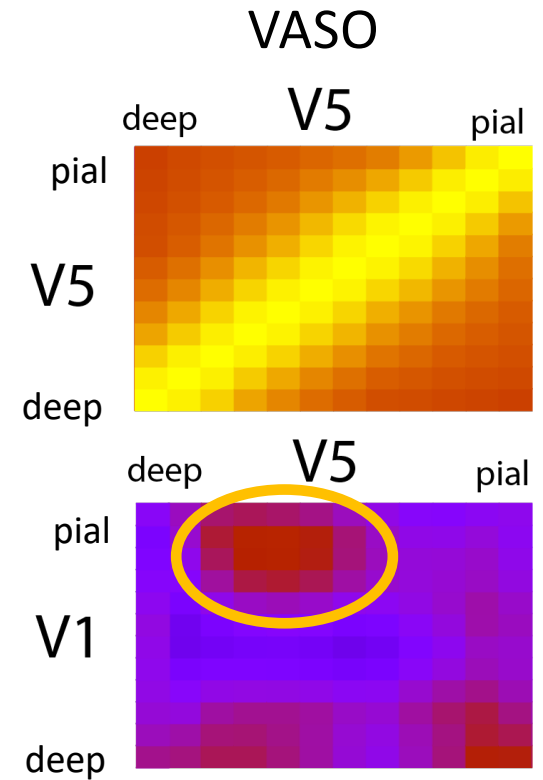
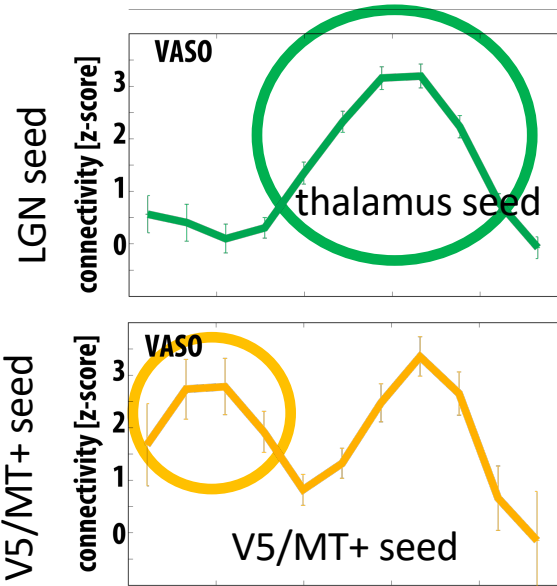
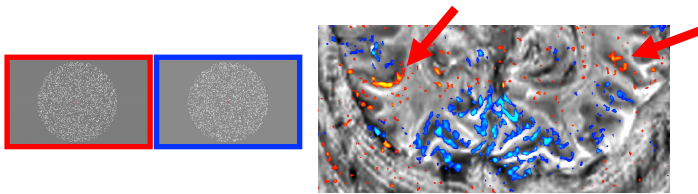
thalamus



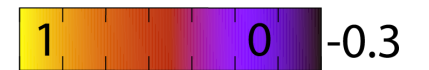
Resting-state connectivity



MT+ (containing V5) is selected based on a functional localizer: motion vs static



Seed-based connectivity in V1 across layers

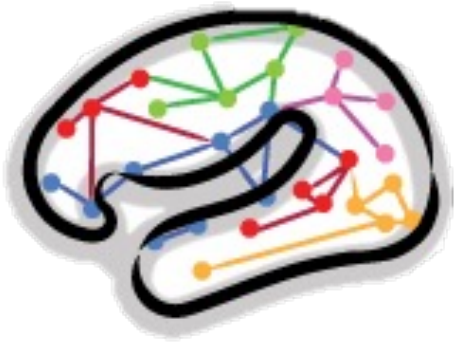


V5-V1 feedback originated in deeper layers of V5 and terminates in upper layers of V1

Free movie watching

Human connectome data:

93 non-sibling HCP participants at 1.6 mm, TR = 1s whole brain
15 min movie consisting of 5 short clips with rest
ICA-fixed, 7T



Participants are clustered in three groups (based on similarity in task positive network).

Layer-fMRI data:

6 participants at 2mm
and 0.8mm (VASO)
TR = 2.5 s, 7T
5 times same movie
HCP signal traces are
taken as regressor
VASO vessel correction

