

Tailored Excitation Pulse Design – Research Update

fMRI Group Meeting

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2024.11.26

Outline

- ISMRM Abstract: *Impact of Spatially Selective Signal Suppression on BOLD fMRI Reliability*
- ROI-Image-Quality Driven RF Pulse Design

Impact of Spatially Selective Signal Suppression on BOLD fMRI Reliability

Submission to ISMRM 2025 Abstract

ROI-Image-Quality Driven RF Pulse Design

Impact of Spatially Selective Signal Suppression on BOLD fMRI Reliability

Submission to ISMRM 2025 Abstract

Motivation: rFOV Imaging

Potential Benefits:

1. Reduced scan time
2. Enhanced image resolution
3. Improved image quality in EPI due to shorten echo train length

Application:

Brainstem/ Spinal cord fMRI ^[1]

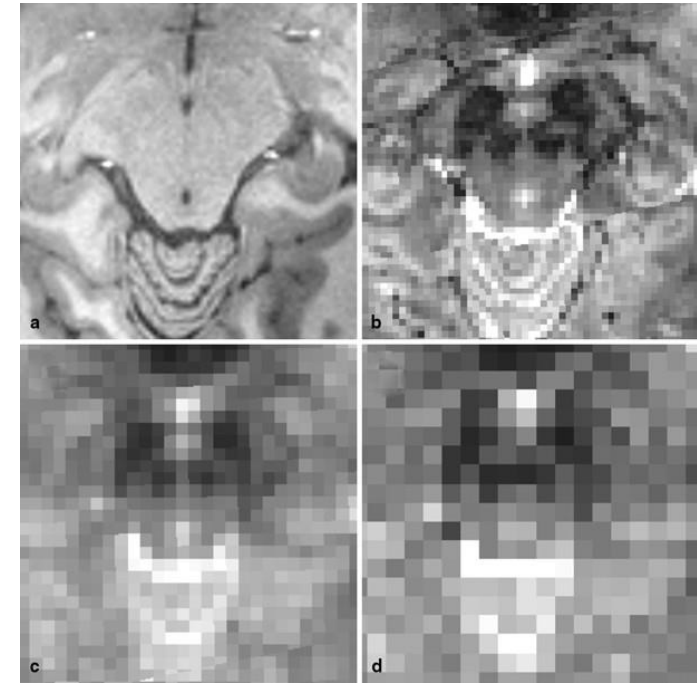


Table 1 Brainstem-specific problems in fMRI and some possible solutions for them

Problem	Possible solutions
Brainstem nuclei are very small compared to cortical structures	<ul style="list-style-type: none">Choose image resolution of ~ 2 mm or higherIncrease SNR by acquiring more volumes, using specialized coils or higher field strengthsUse parallel imaging, multi-band imaging or partial FOV acquisitions to minimize spatial distortions and keep scanning time within limitsUse spatial smoothing with caution (≤ 3 mm FWHM)

[1]Beissner, F., *Clin Neuroradiol* (2015) 25:251–257

Motivation: rFOV Imaging

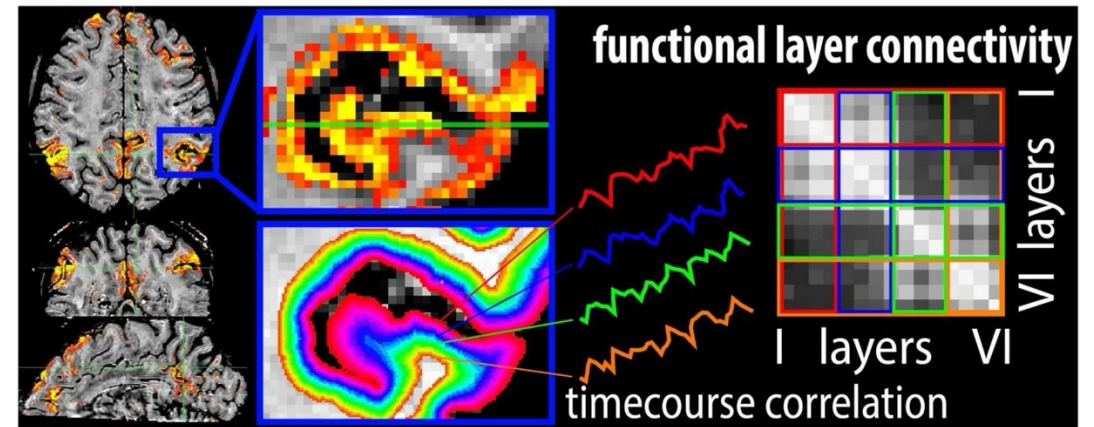
Potential Benefits:

1. Reduced scan time
2. Enhanced image resolution
3. Improved image quality in EPI due to shorten echo train length

Application:

Brainstem/ Spinal cord fMRI ^[1]

Layer-specific fMRI ^[2]



motor cortex), the approach of thicker MRI-slices can help improve sensitivity. However, to fully grasp the neural representation of laminar activation across the folded cortical ribbon in higher-order brain areas with variable folding patterns, isotropic submillimeter resolutions are vital.

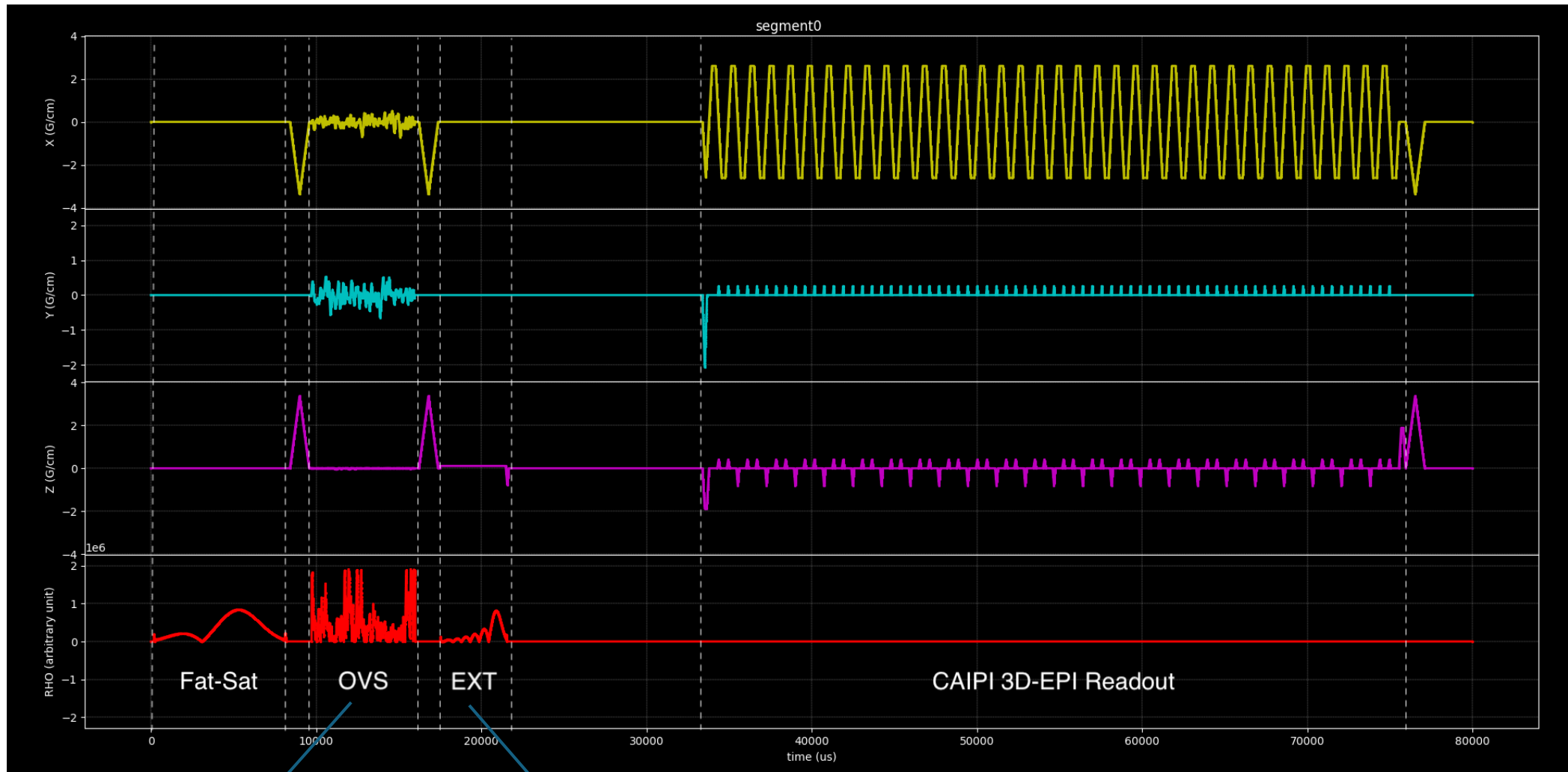
[1]Beissner, F., *Clin Neuroradiol* (2015) 25:251–257

[2]Huber, L., et al., *Progress in Neurobiology* 207 (2021) 101835

Synopsis

- We can restrict FOV by a short (6ms) outer-volume suppression (OVS)
- However, the impact of OVS on the sensitivity/specificity of fMRI detection is unclear
- We obtained 7.5-fold accelerated **CAIPI 3D-EPI** BOLD fMRI data during four repetitions of a block finger-tapping task in a healthy subject, both **w/ and w/o OVS**. We assessed test-retest reliability of the activation maps using **receiver-operating-characteristic** (ROC) analysis.

Sequence: SPGR with CAIPI 3D-EPI readout



2D in-plane selective;
6ms

1D slab-selective;
2ms

OVS: Outer-volume suppression

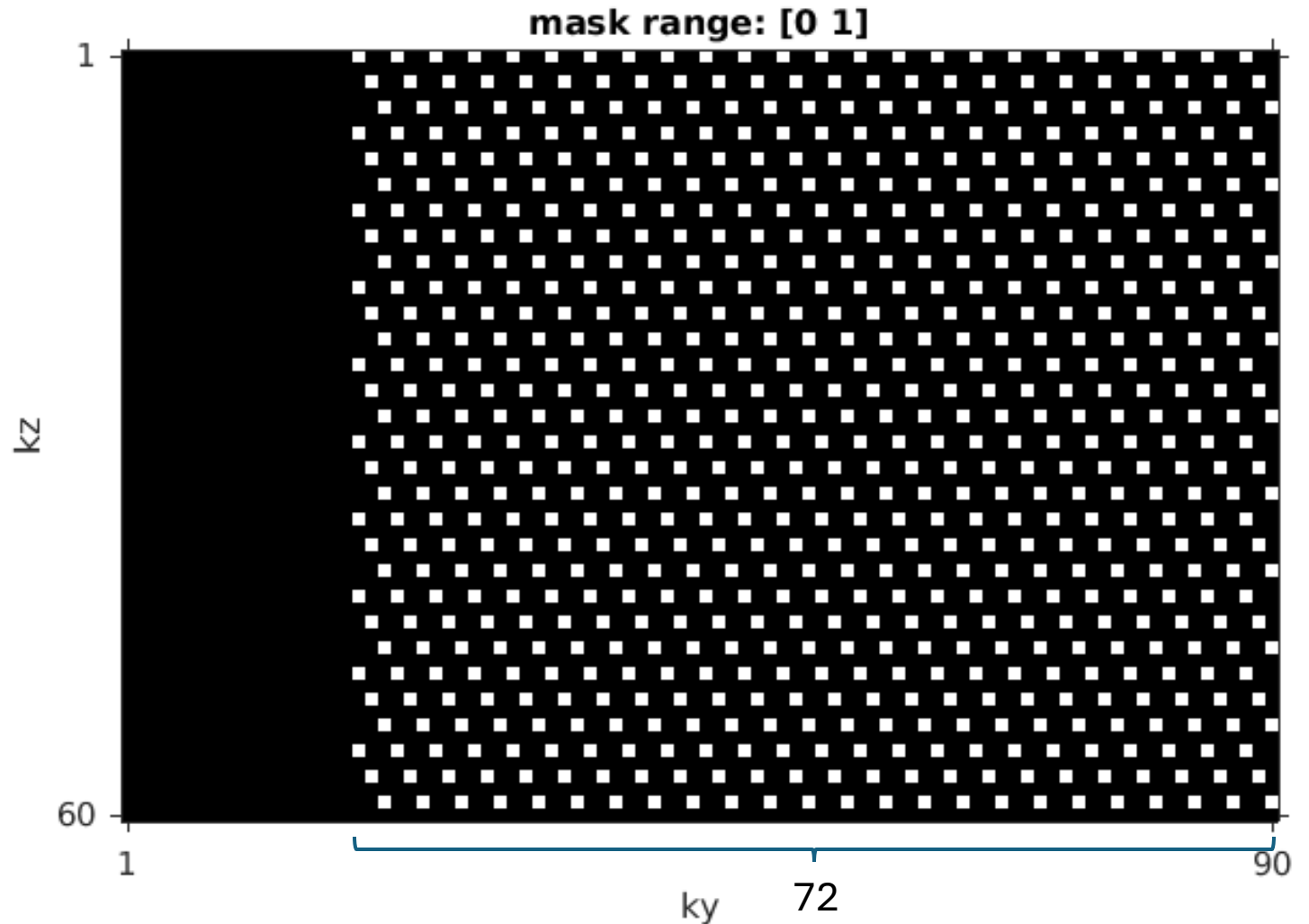
CAIPI Sampling Pattern

Ry= 3; Rz=2
Partial ky = 72/90

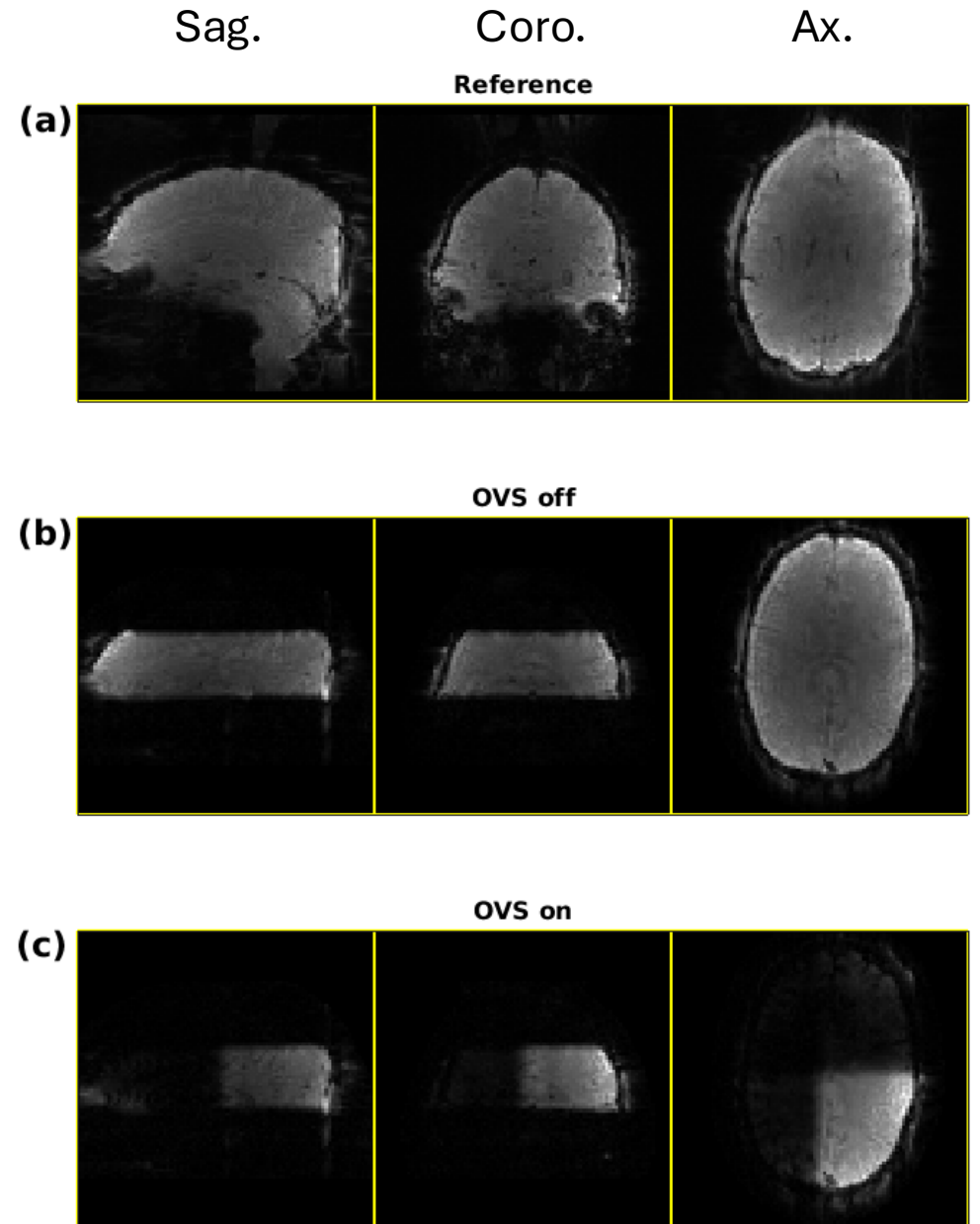
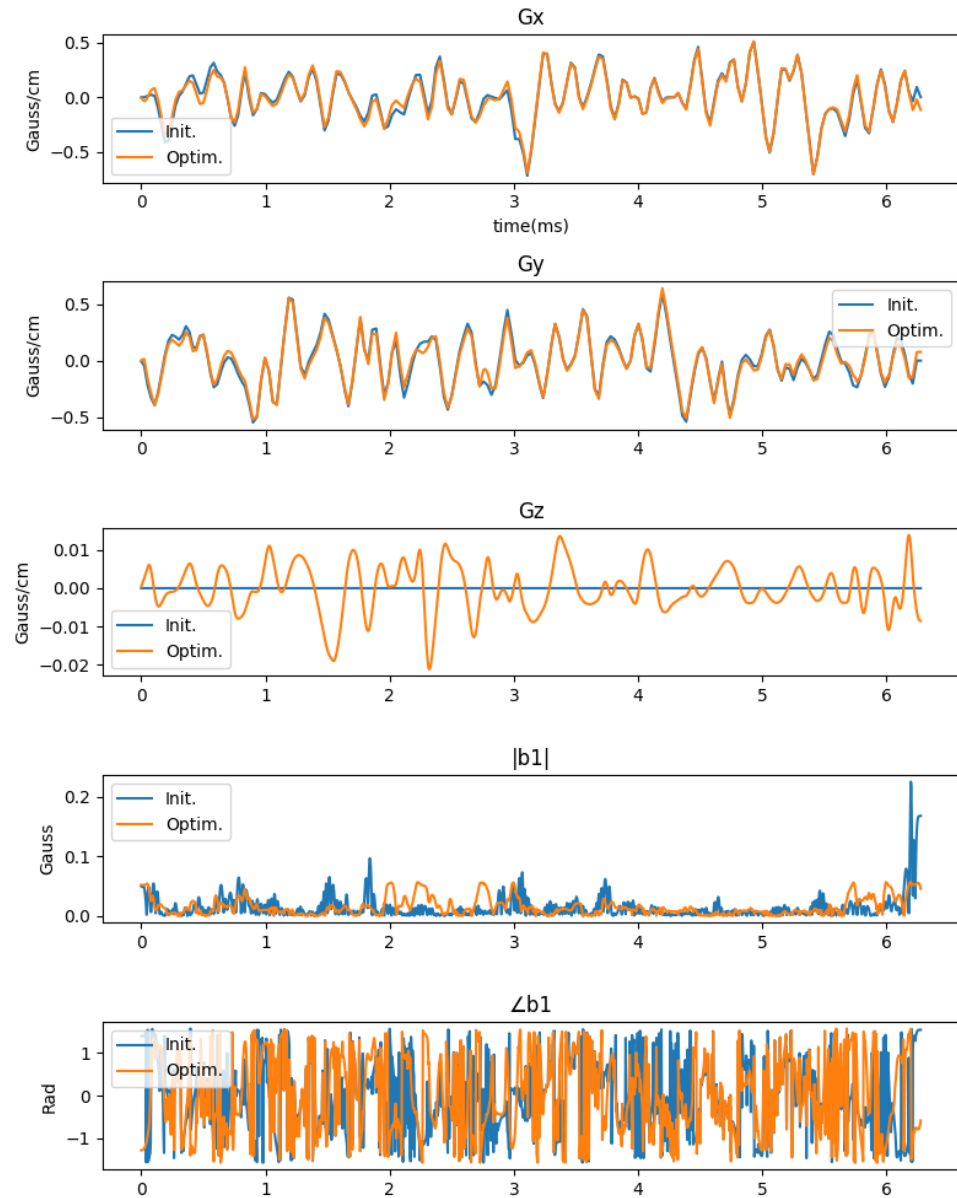
Matrix size:
90x90x60

Resolution:
2.4mm³

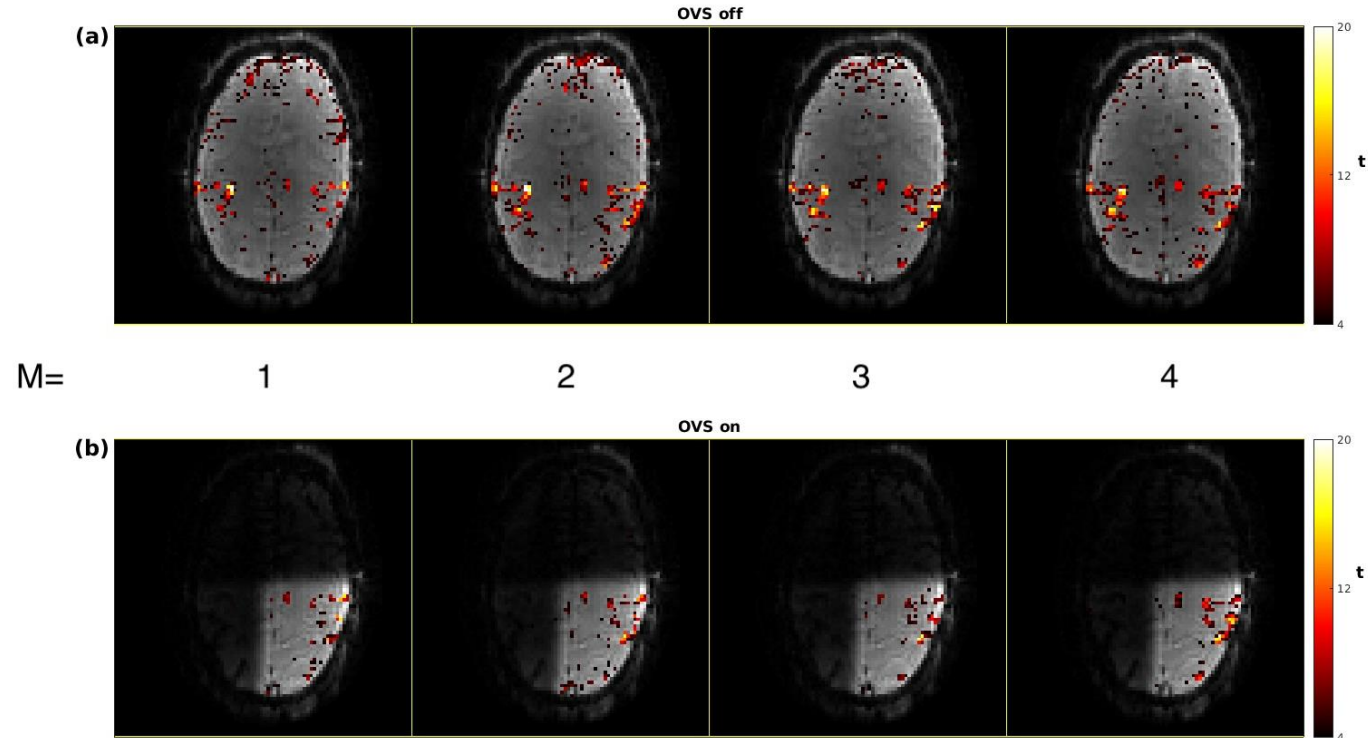
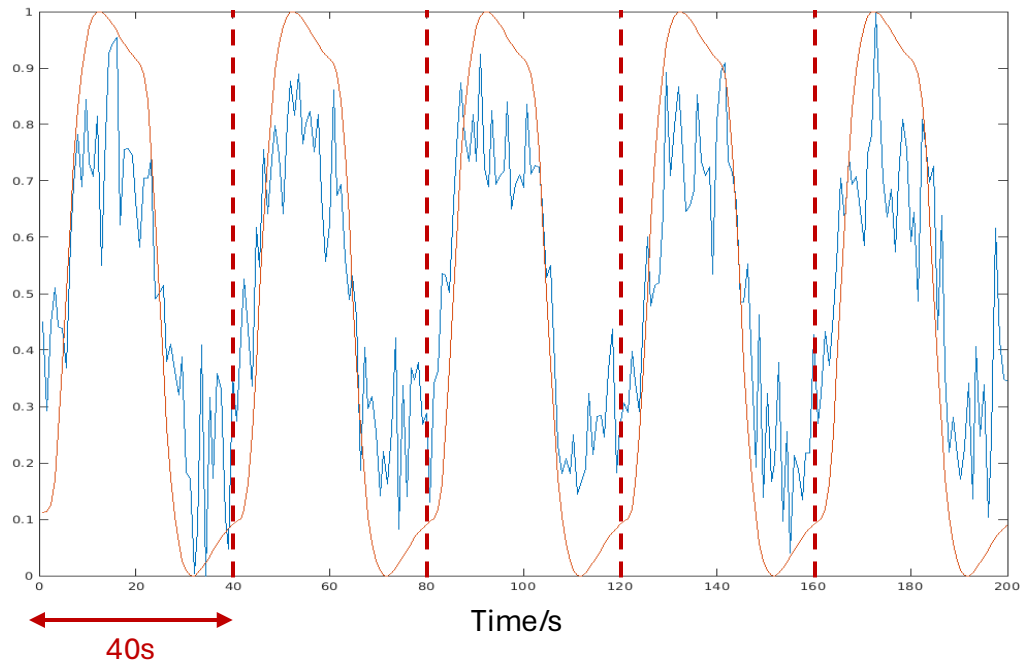
TR: 0.8s
TE=30ms



Structural Scan



fMRI Finger-tapping test

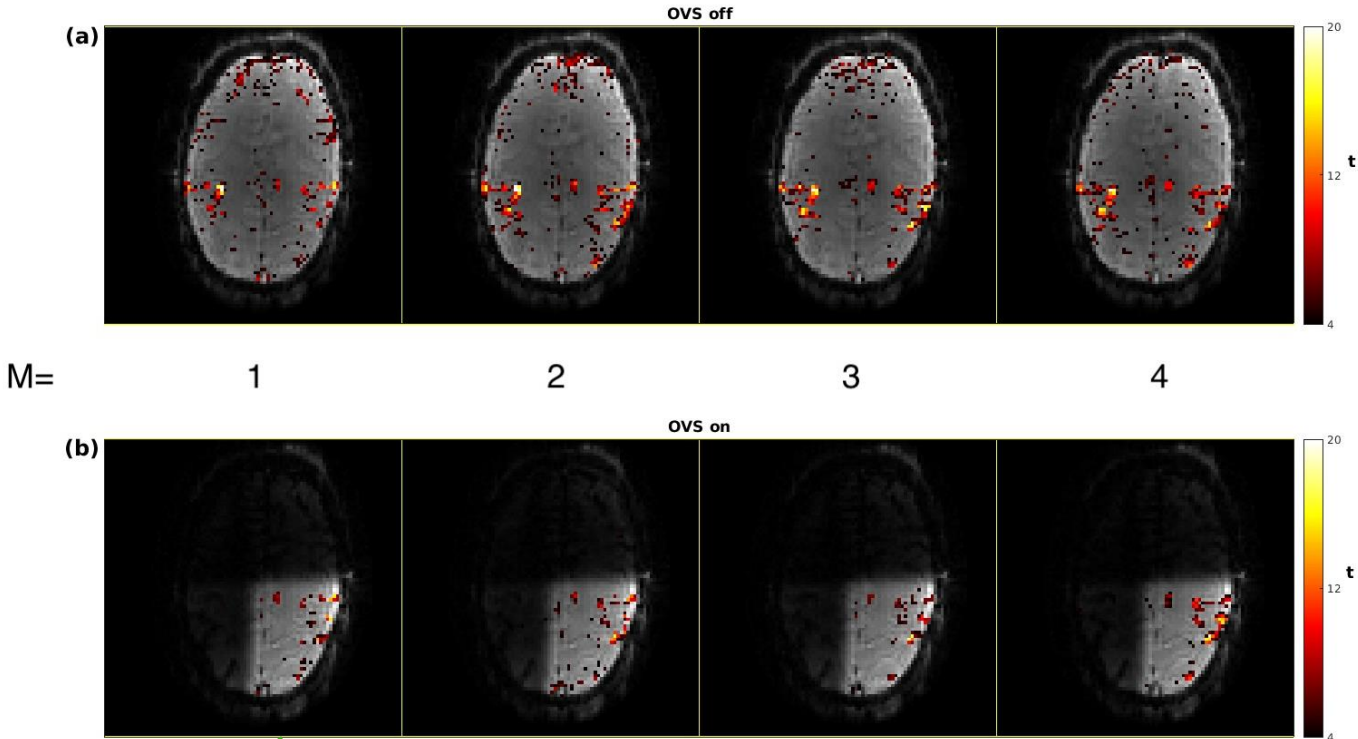
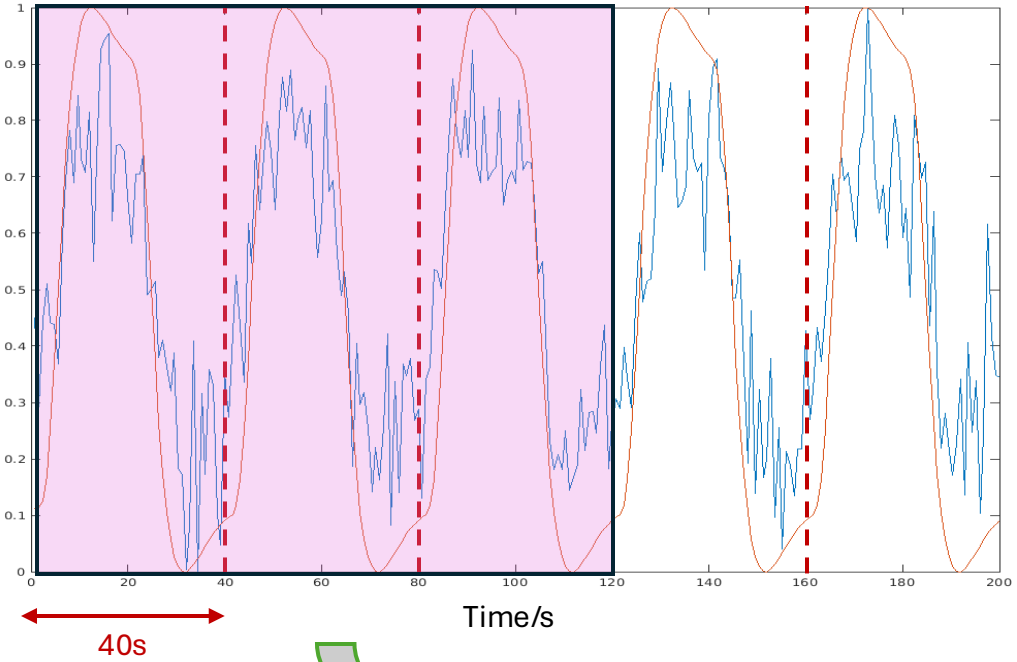


20s rest + 20s tapping = 1 block

5 blocks = 1 repetition

4 repetitions for both OVS off and on, alternatively on-off-on-off-on-off-on-off

fMRI Finger-tapping test



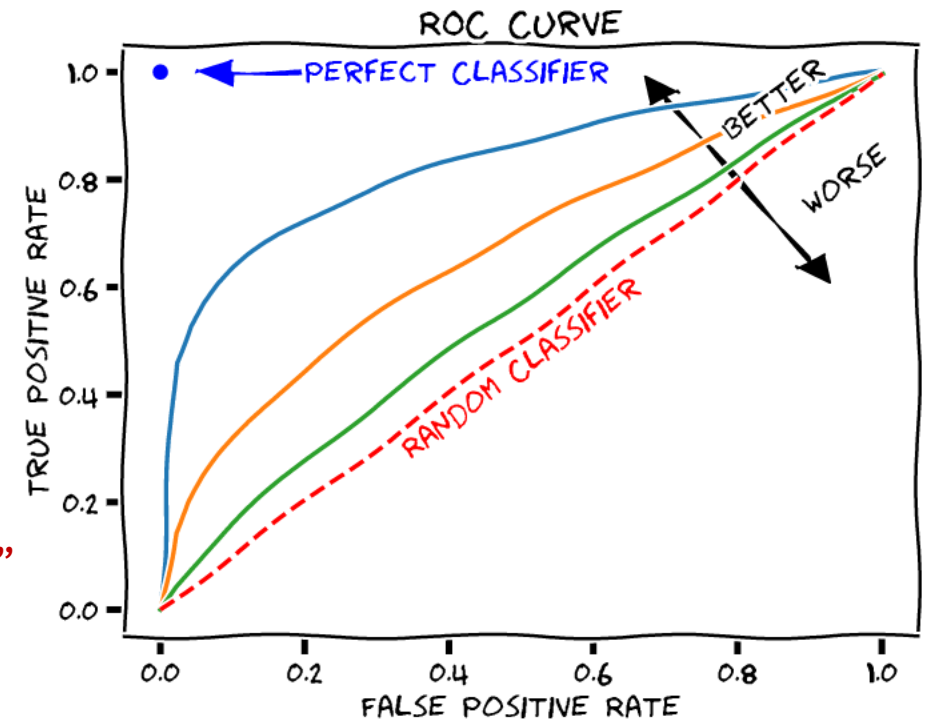
T-score maps; Threshold at 4

Test-Retest Reliability

Sweep the threshold value, for each threshold, calculate (FPR,TPR), get a dot in the ROC curve

$$\text{FPR} = \frac{\text{FalsePositive}}{\text{FalsePositive} + \text{TrueNegative}} \triangleq p_I \text{ "False Alarm"}$$

$$\text{TPR} = \frac{\text{TruePositive}}{\text{FalseNegative} + \text{TruePositive}} \triangleq p_A \text{ "Hit"}$$

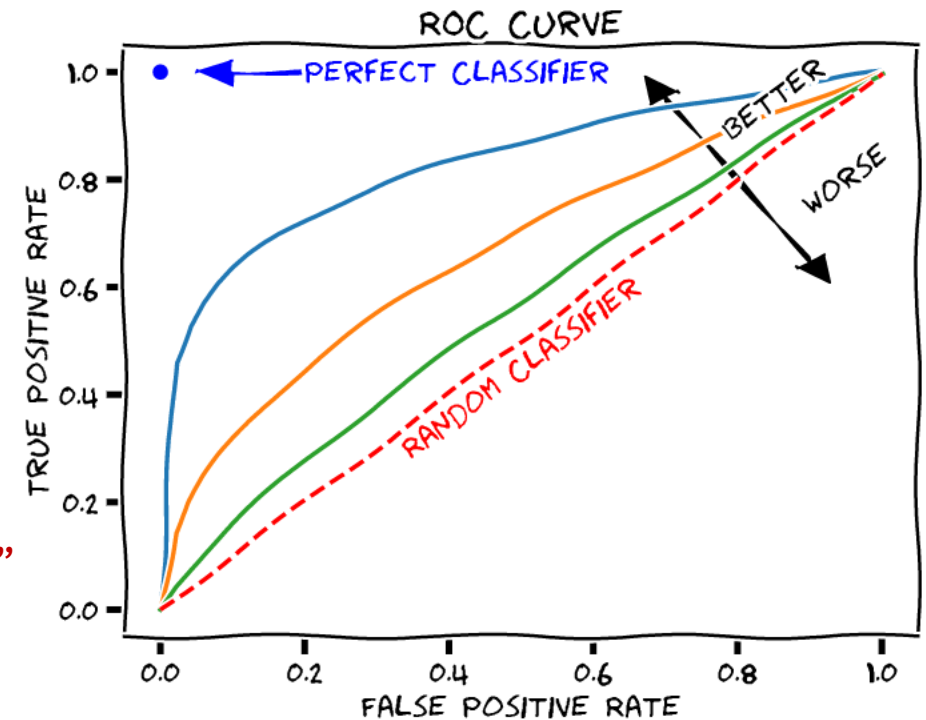


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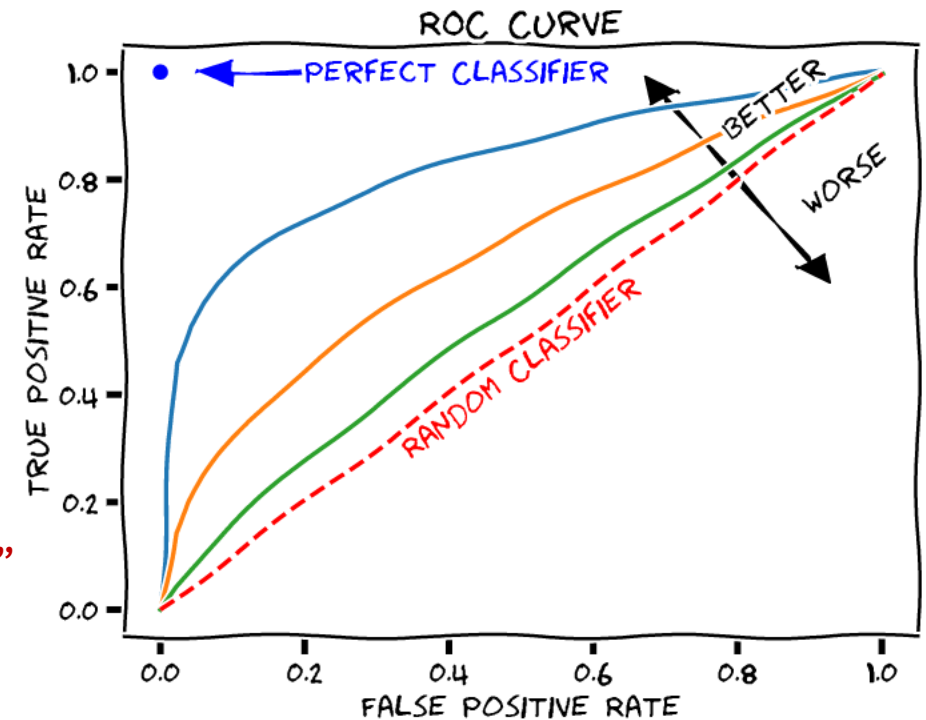
**Q: Which one is better: guess all positive, or guess all negative?
(suppose in fMRI, proportion of truly activated voxels is far less than 1/2)**

Test-Retest Reliability

Sweep the threshold value, for each threshold, calculate (FPR,TPR), get a dot in the ROC curve

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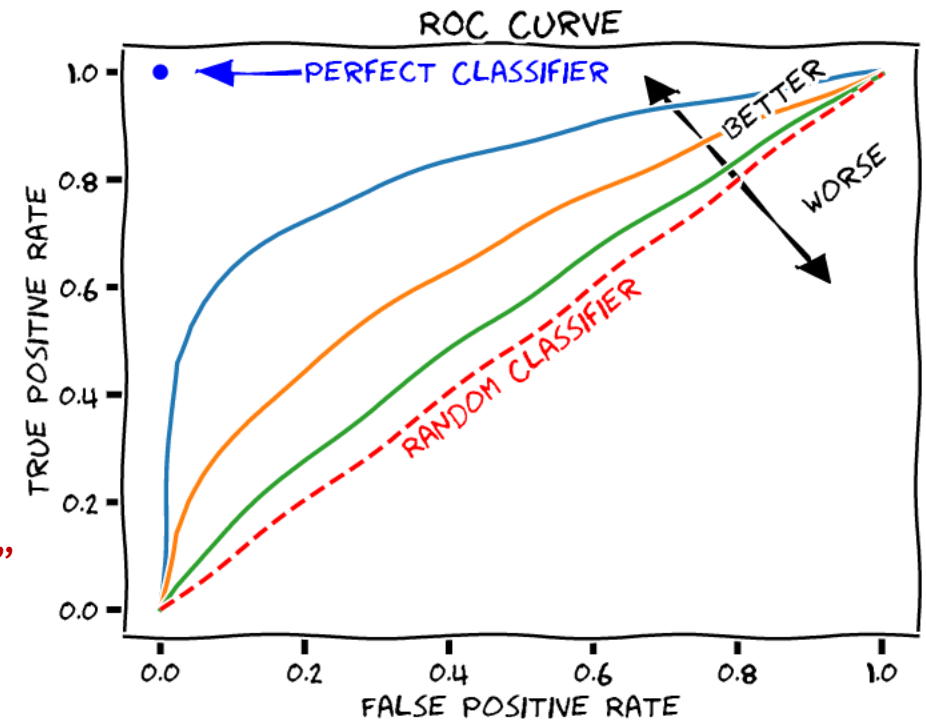
Q: How do we know the ground truth activation classification?

Test-Retest Reliability

Sweep the threshold value, for each threshold, calculate (FPR,TPR), get a dot in the ROC curve

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Q: How do we know the ground truth activation classification?

- Long scan;
- **Statistic Model**

Entails multiple repetitions ($M \geq 4$) of experiment

Mixed-Binomial Model^[1,2]

Assumptions:

- The behavior of voxels across different trials is i.i.d.
- The behavior of each voxel is independent of other voxels
- All voxels behave according to the same probability distribution

[1] Genovese, C.R, et al.,(1997). *Magn. Reson. Med.*, 38: 497-507.

[2] Noll, D.C., et al., (1997). *Magn. Reson. Med.*, 38: 508-517.

Mixed-Binomial Model

λ : proportion of truly active voxels

p_A : TPR

p_I : FPR

Raw reliability map:

R_v = Number of times out of M repetitions a voxel v is classified active

Assume R_v is drawn from a mixture of two binomial distributions:

$$R_v \sim \lambda \cdot \text{Binomial}(M, p_A) + (1 - \lambda) \cdot \text{Binomial}(M, p_I)$$

Due to independence assumption, the likelihood function of parameters

p_A, p_I, λ , only depends on the counts:

$$n_k = \sum_{v \in V} \mathbb{I}_{\{R_v=k\}} = \text{Number of voxels that are classified active } k \text{ out of } M \text{ repetitions}$$

Mixed-Binomial Model

Let $\mathbf{n} = (n_0, n_1, \dots, n_M)$ be the **histogram-vector**.

(The log of the) Posterior likelihood function of parameters p_A, p_I, λ :

$$l(p_A, p_I, \lambda | \mathbf{n}) = \ln \mathbb{P}((p_A, p_I, \lambda | \mathbf{n})) \cong \sum_{k=0}^M n_k \ln[\lambda p_A^k (1 - p_A)^{(M-k)} + (1 - \lambda) p_I^k (1 - p_I)^{(M-k)}]$$

We estimate the parameters by the method of **Maximum Likelihood (ML)**.

Mixed-Binomial Model

Dependent likelihood model:

We use the same statistic maps (e.g., t-score) to generate a series of reliability maps by selecting K different thresholds $\tau_0 < \tau_1 < \dots < \tau_{K-1}$.

These reliability maps should **share a common λ** , while at each τ_k , the points $(p_I^{(k)}, p_A^{(k)})$ are different, $k = 0, \dots, K - 1$.

Define: p_{AK} is the probability that a truly active voxel is classified active at k of the threshold levels and similarly for p_{IK} , $k = 0, \dots, K$.

Note that $p_{AK} = \sum_{j=0}^{K-1} p_{Aj}$, $p_{IK} = \sum_{j=0}^{K-1} p_{Ij}$.

Mixed-Binomial Model

Dependent likelihood model:

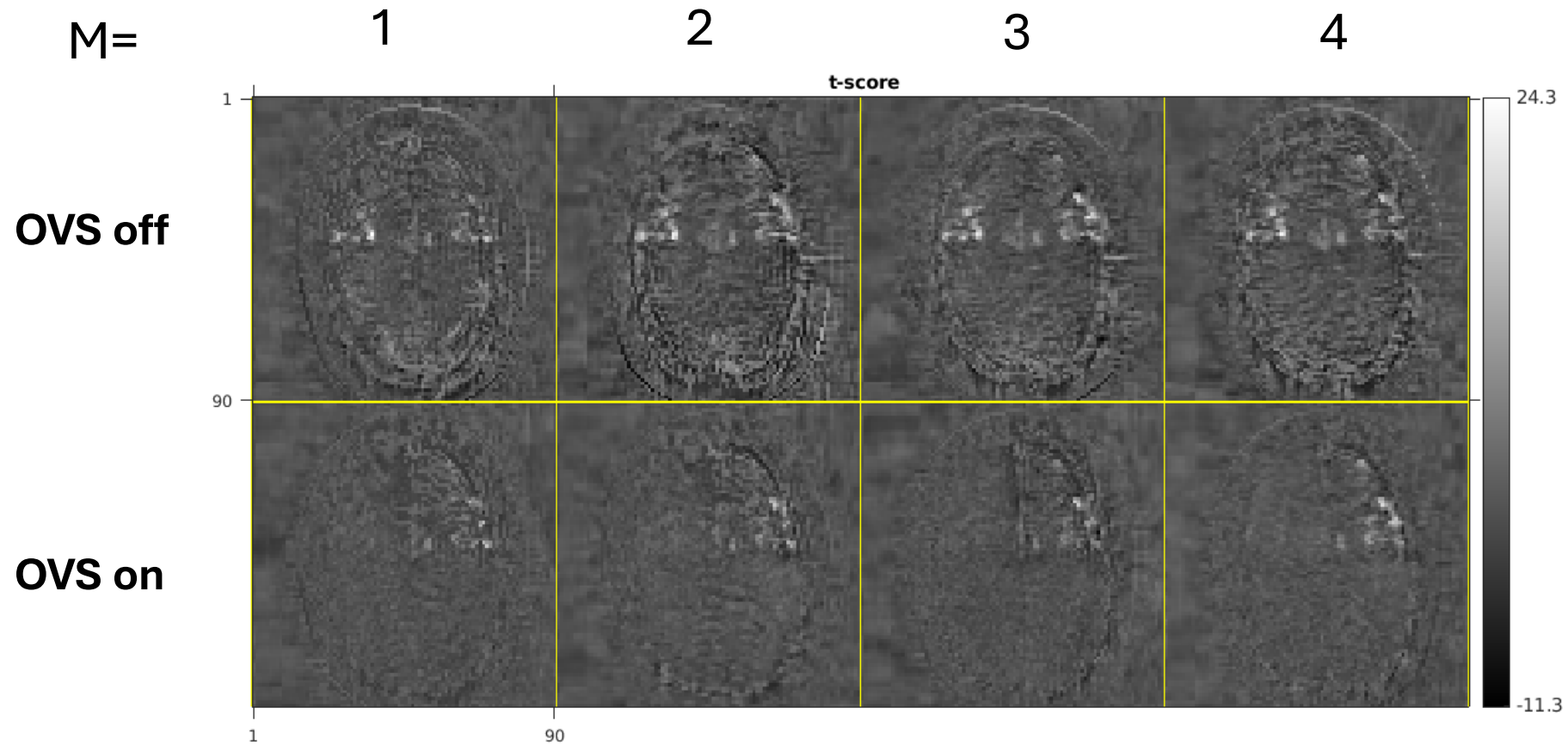
Define $n_{\mathbf{t}}$ for $\mathbf{t} = (t_0, \dots, t_K)$, to be number of voxels classified active at k threshold levels t_k times (out of M) for each $k = 0, \dots, K$.

The dependent likelihood function is:

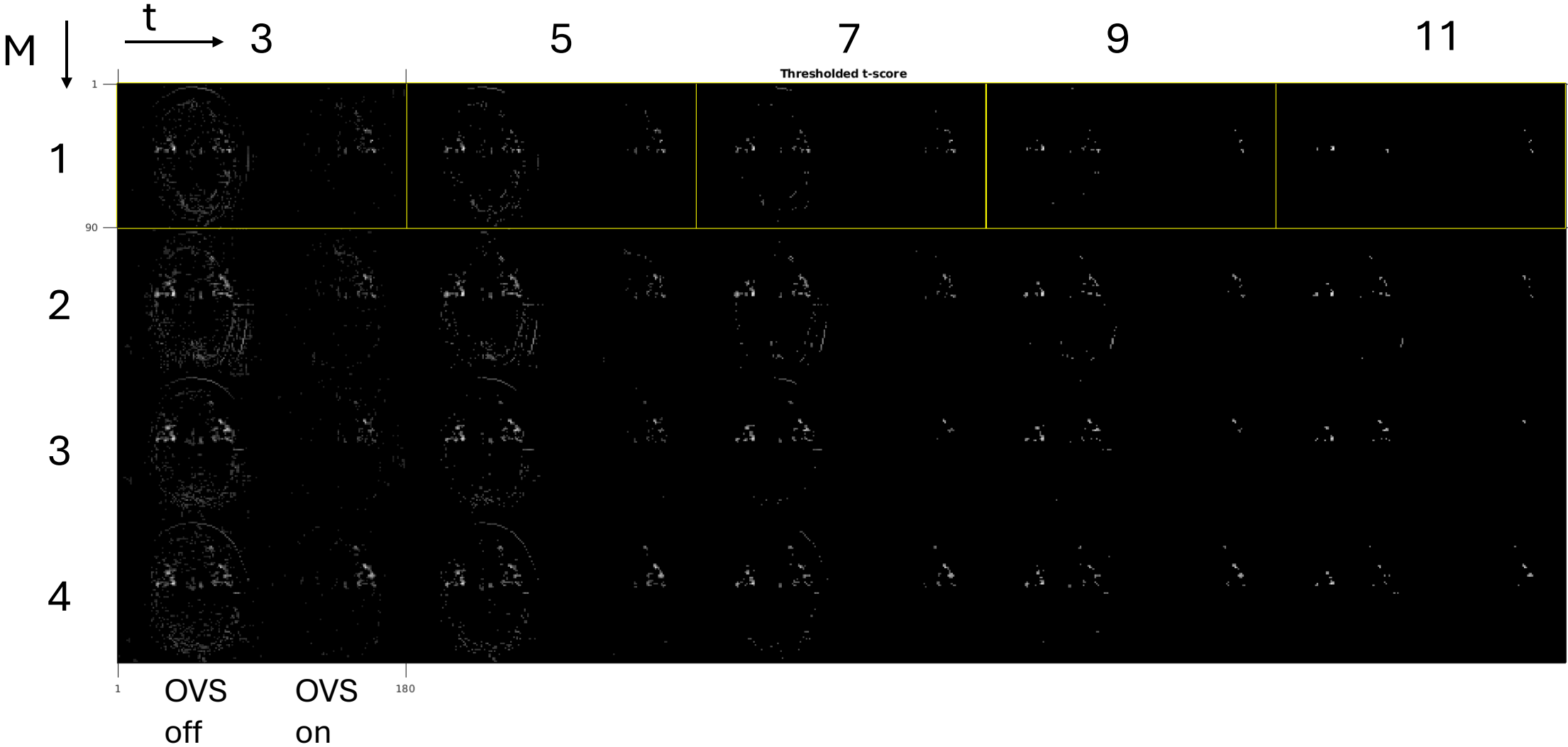
$$l_{dep}(\mathbf{p}_A, \mathbf{p}_I, \lambda \mid \mathbf{n}) = \sum_{\mathbf{t}} n_{\mathbf{t}} \ln \left[\lambda \prod_{k=0}^K p_{Ak}^{t_k} + (1 - \lambda) \prod_{k=0}^K p_{Ik}^{t_k} \right]$$

The parameters of interest: $p_A^{(k)} = \sum_{j=k}^K p_{Aj}$, $p_I^{(k)} = \sum_{j=k}^K p_{Ij}$

Raw t-score maps

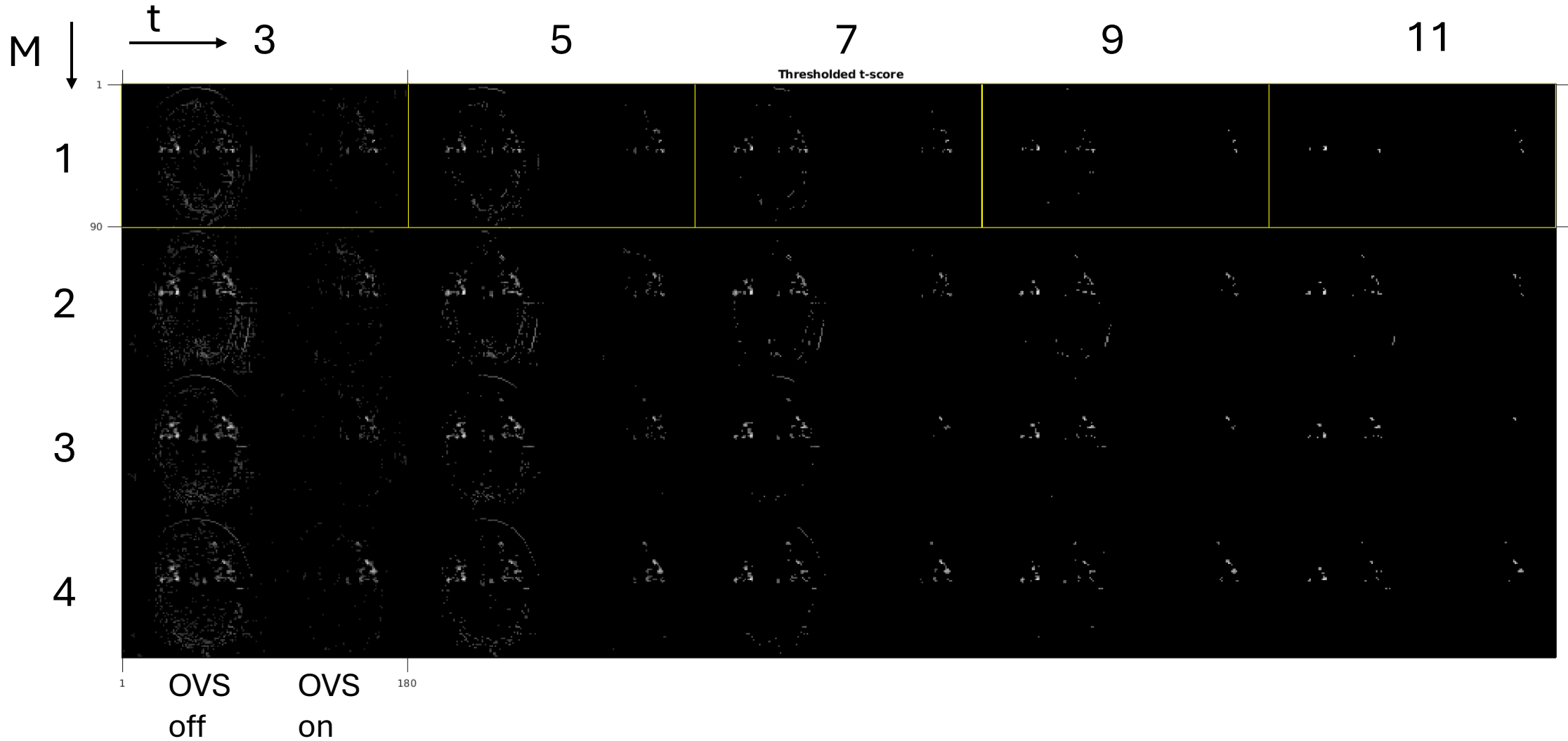


Thresholded t-score maps

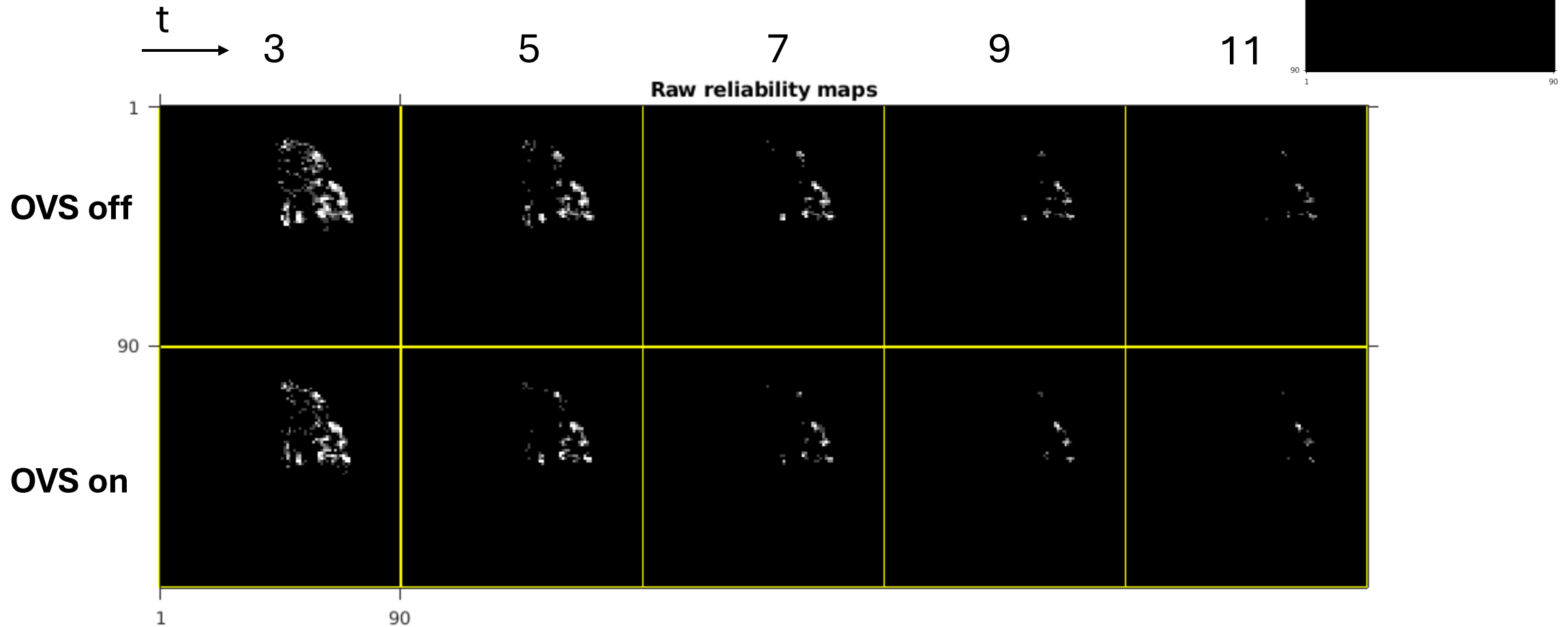


Thresholded t-score maps

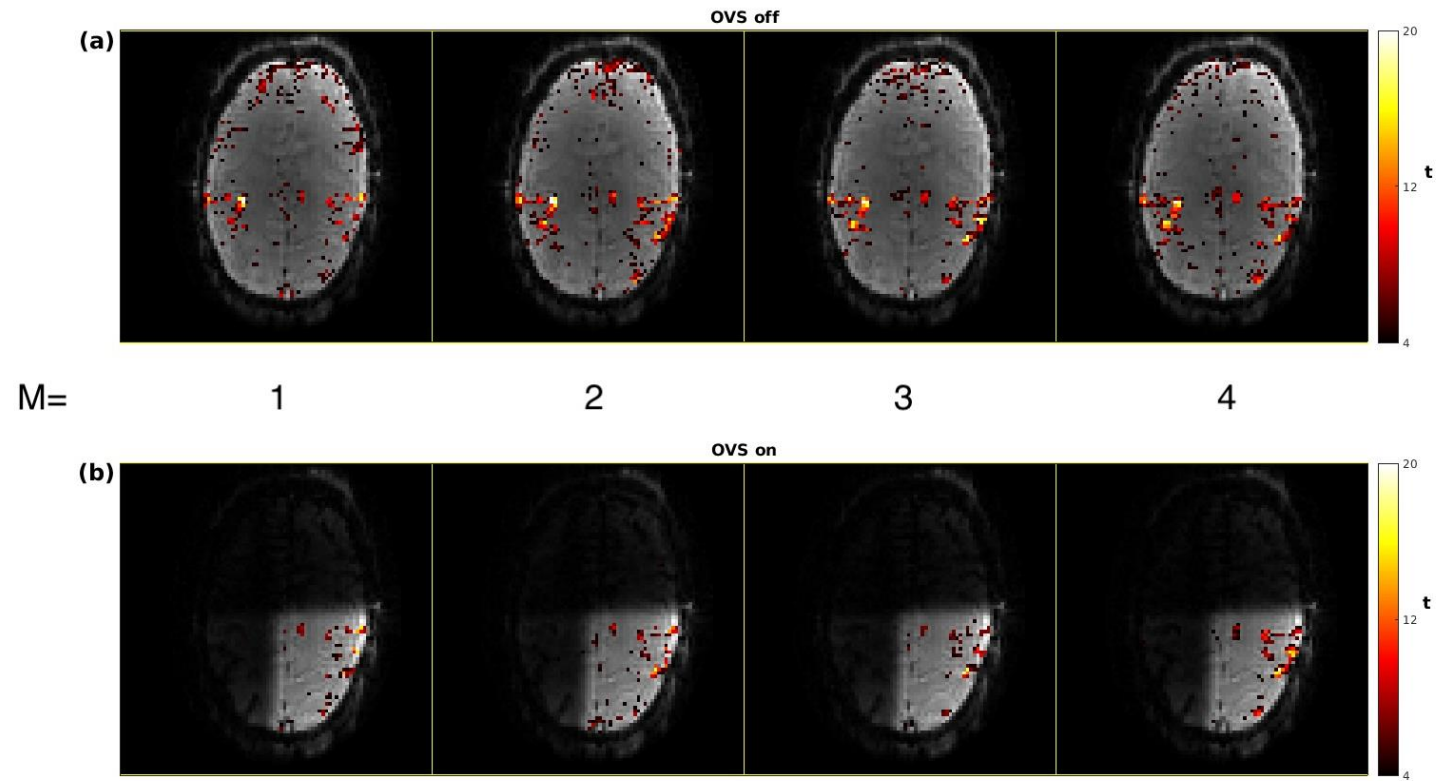
Q: Suppose a voxel is classified active at $k=3$ of the thresholds, which three would it be?



Masked Reliability Maps



Activation Maps



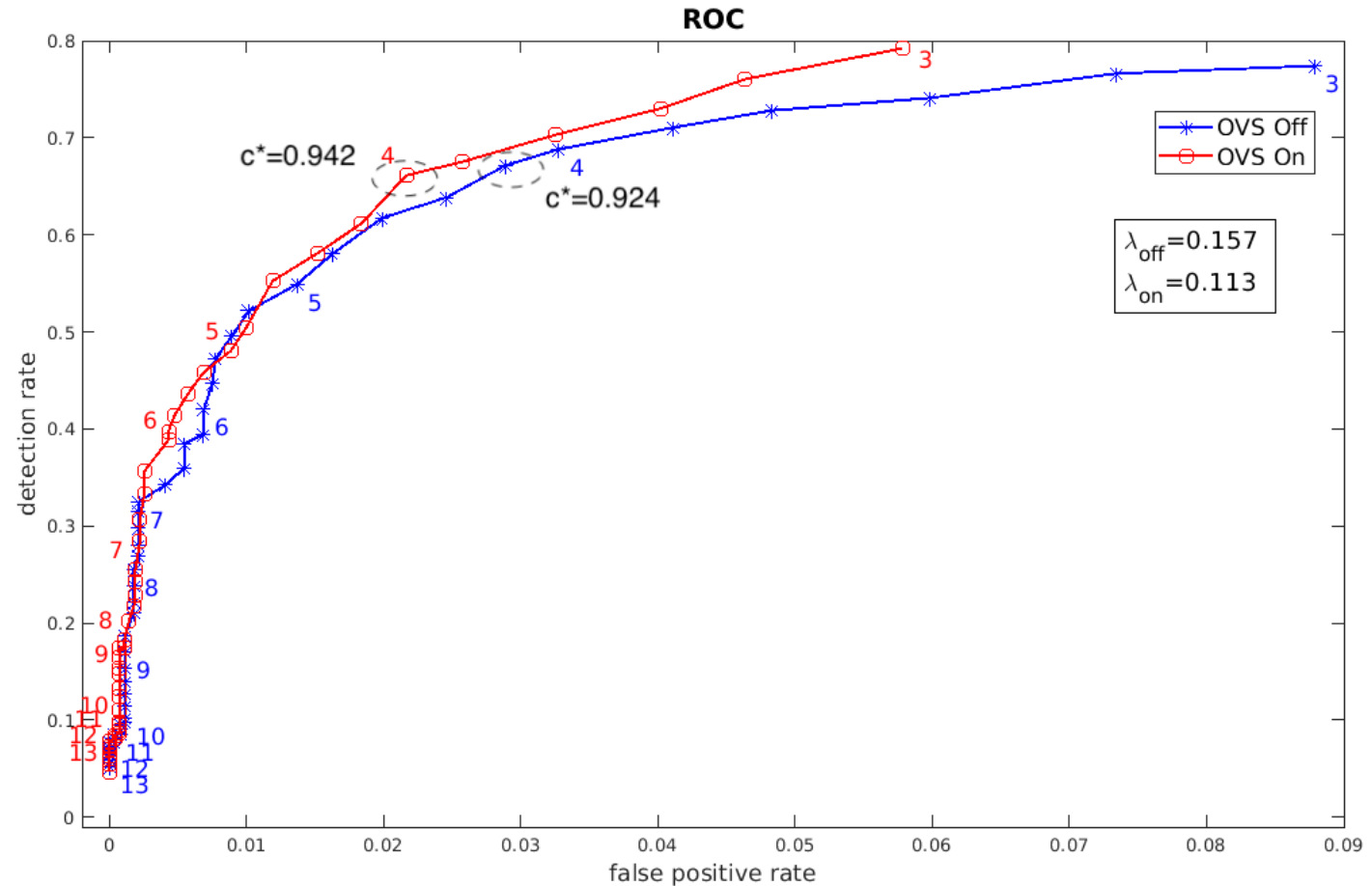
Receiver operating characteristic (ROC) curve

Optimal threshold:

$$\tau^* = \arg \max c(\tau) = \lambda p_A + (1 - \lambda)(1 - p_I)$$

Equivalently, at optimal threshold, the local slope of the curve is $(1 - \lambda)/\lambda$.

At this point on the curve, the cost of removing one false positive voxel is the loss of one truly activating voxel.



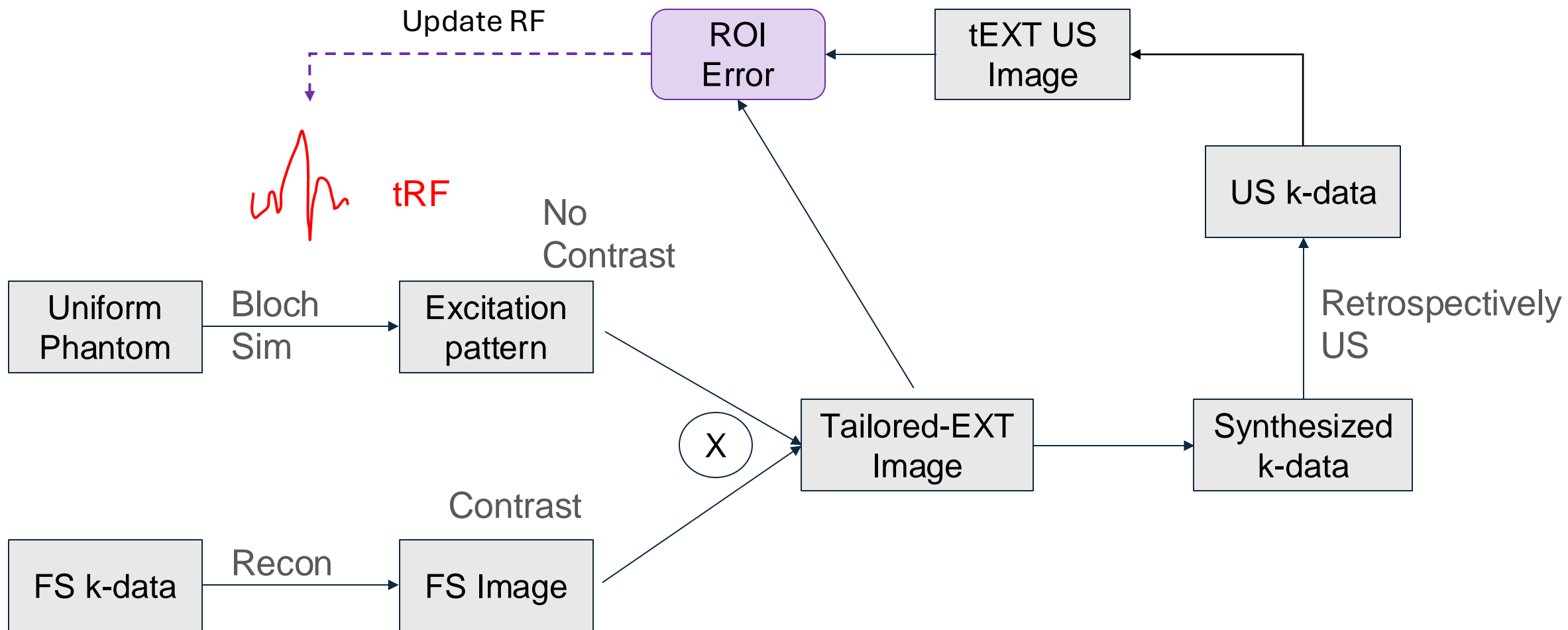
Conclusion

- We demonstrated in a motor task that our OVS pulse preserves sensitivity to BOLD fMRI activation.
- We also show that OVS pulse has somewhat improved test-retest reliability, though further investigation is needed for generalizability.
- We speculate the slight t-score decrease in OVS-on might be caused by the suppression of some in-flow spins from OV which may reduce the BOLD signal.

ROI-Image-Quality Driven RF Pulse Design

Framework

FS: Fully-sampled
US: Under-sampled
tRF: Tailored RF
tEXT: Tailored Excitation



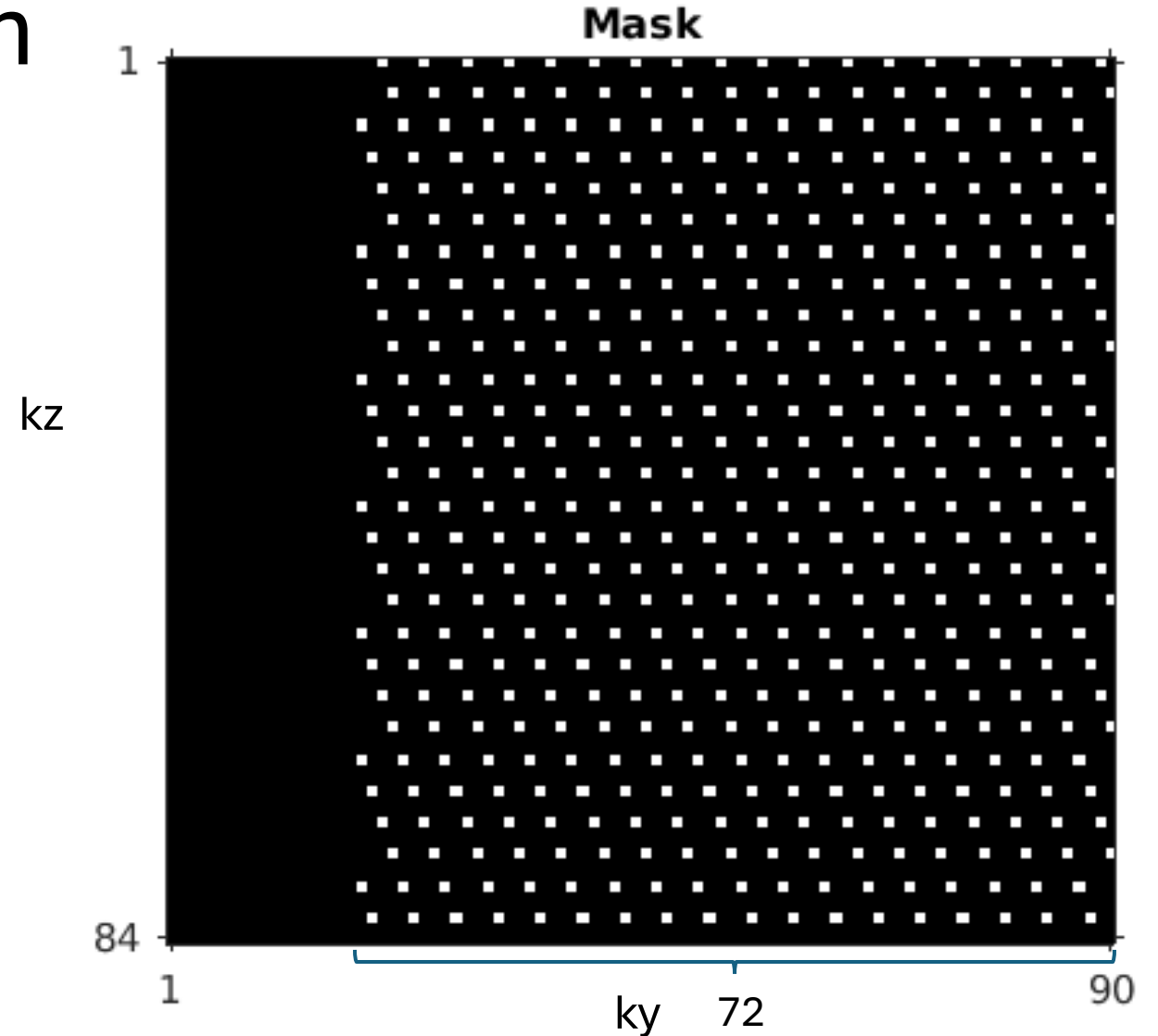
CAIPI Sampling Pattern

$R_y = 4; R_z = 3$

Partial $k_y = 72/90$

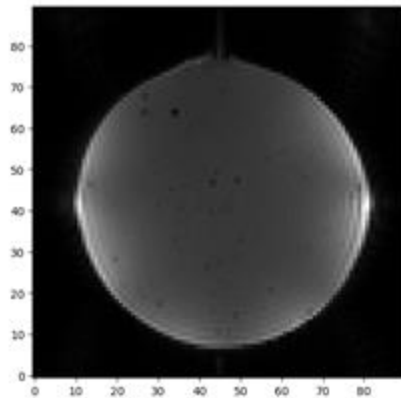
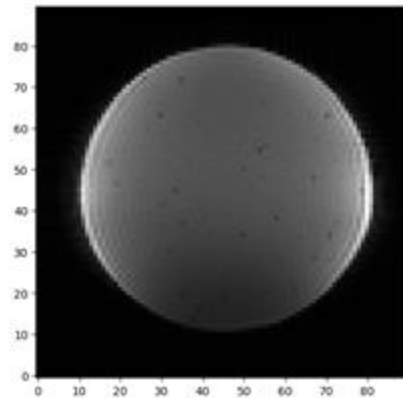
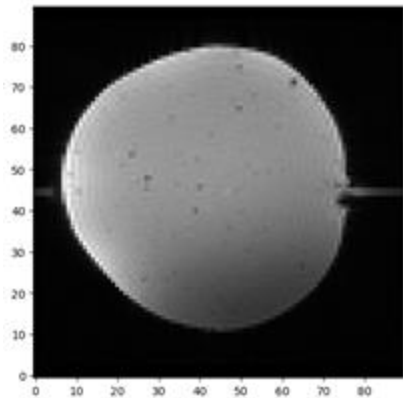
Matrix Size:

$90 \times 90 \times 84$



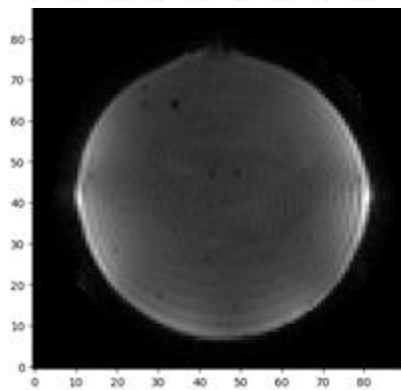
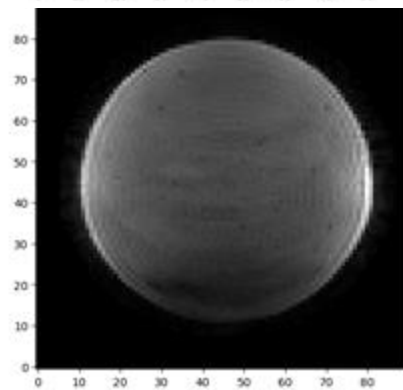
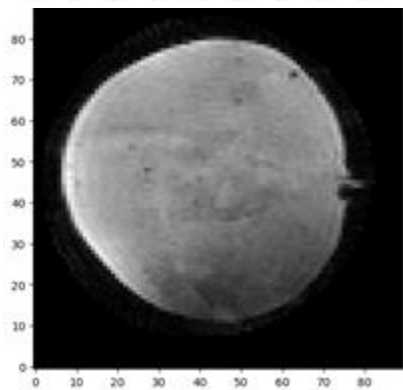
Sanity Check

Fully-sample



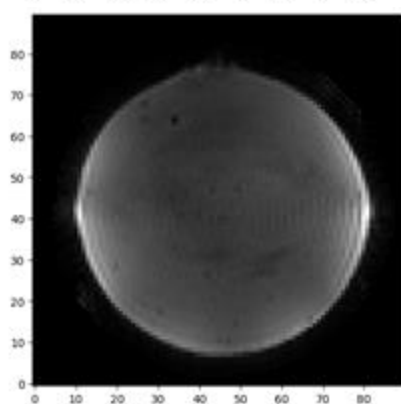
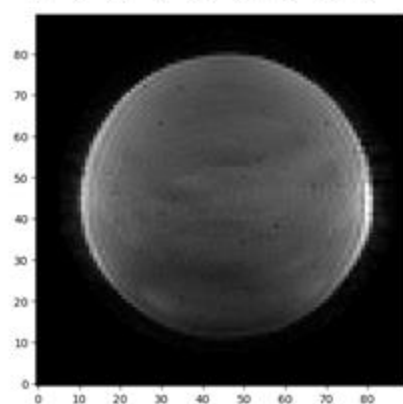
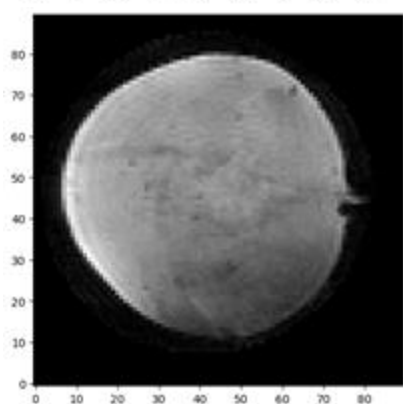
R=1

Retrospective



R=12;
(pky=72/90)

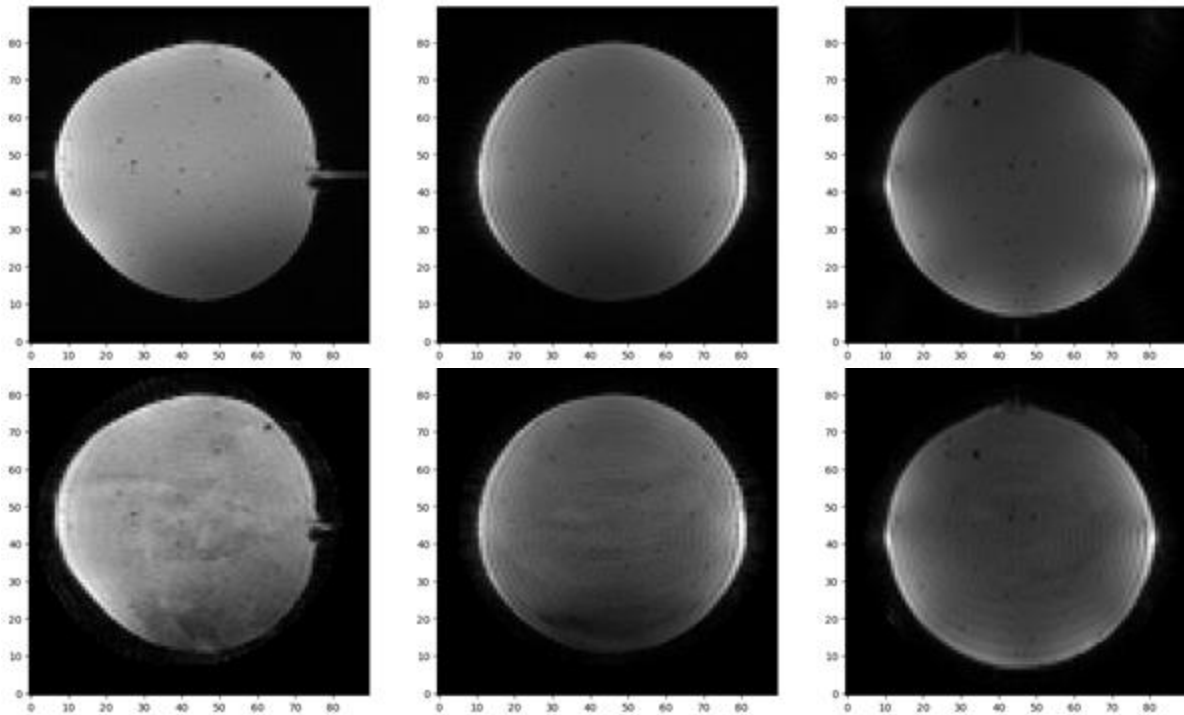
Prospective



R=12;
(pky=72/90)

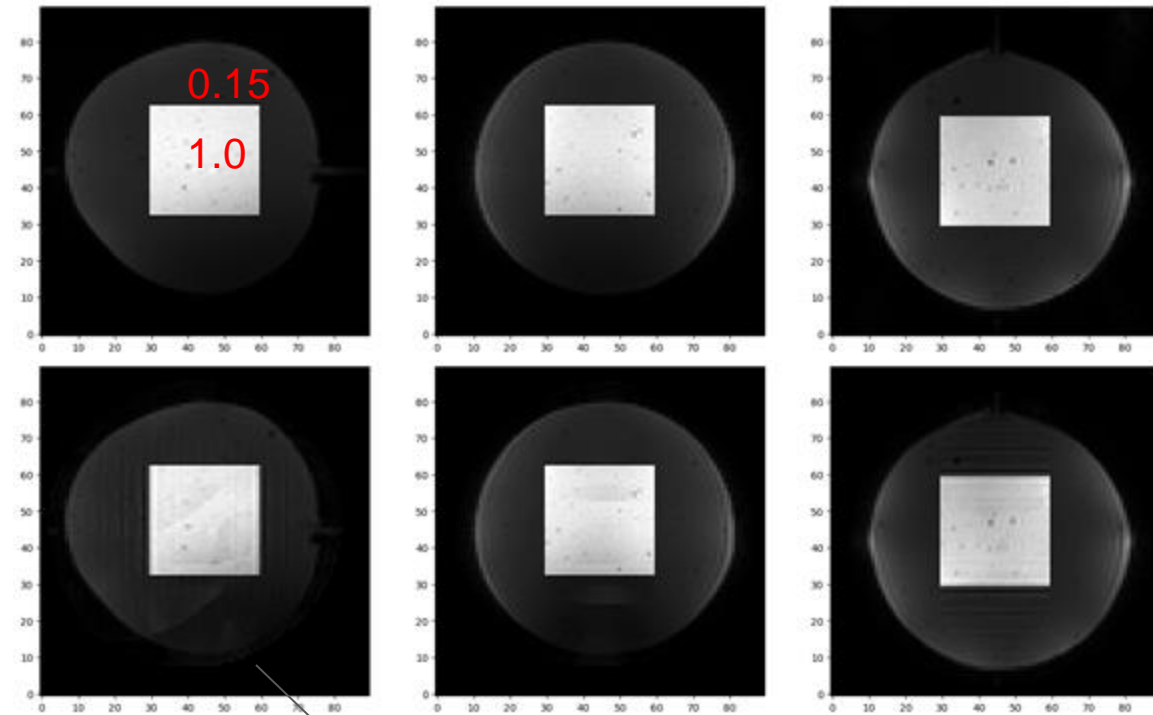
Artificial Weighting to Emulate tailored EXT

Fully-sampled



Retrospective
under-sampled
 $R=12$

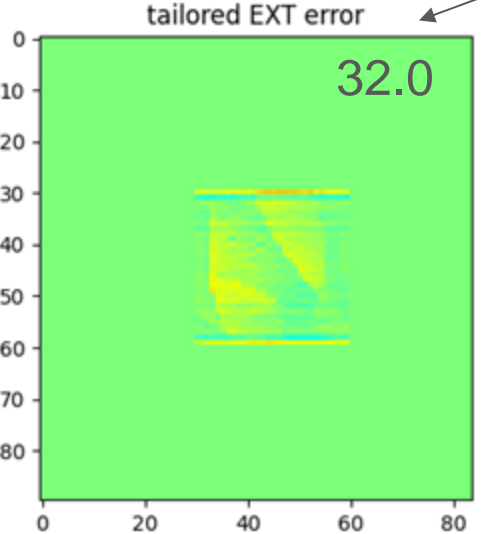
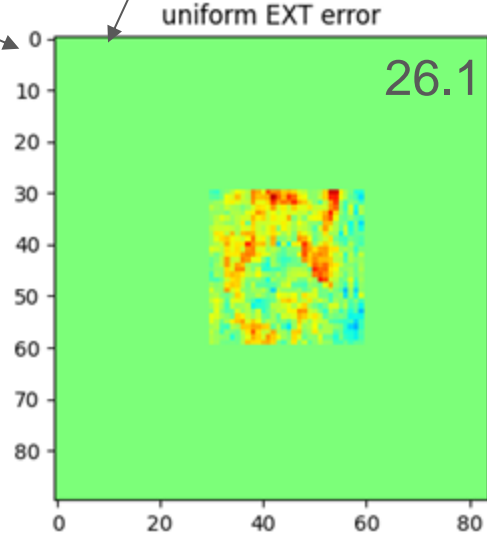
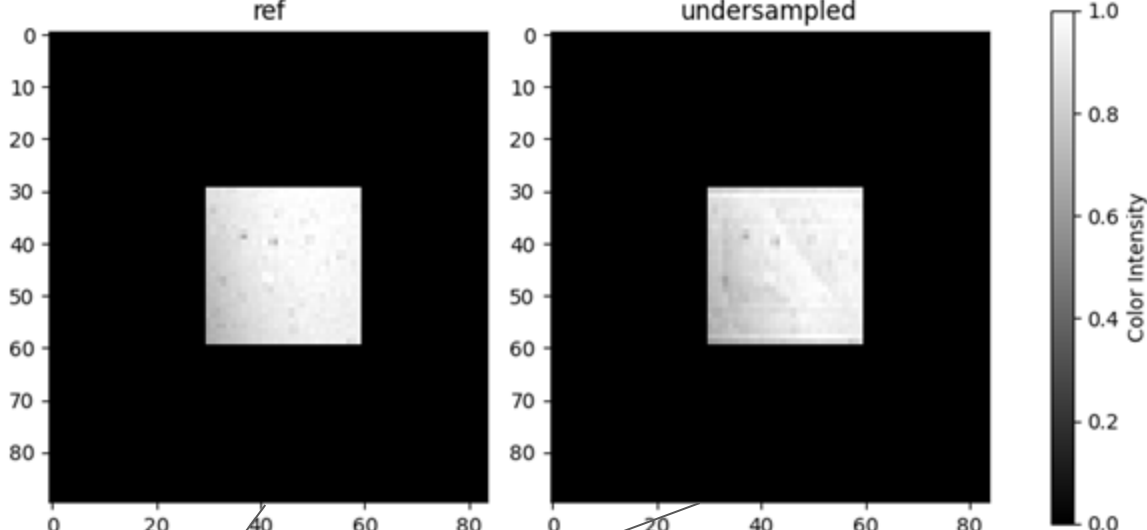
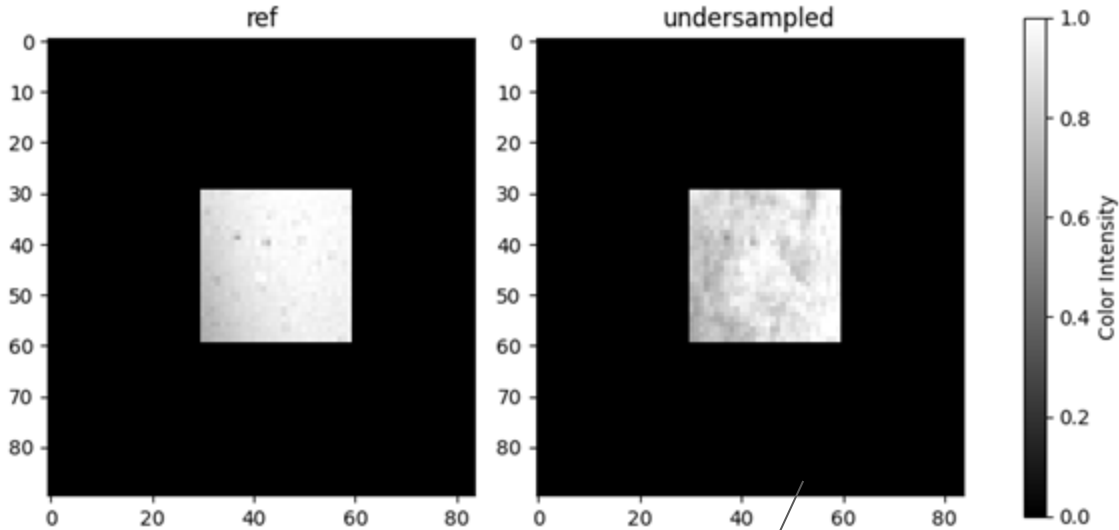
Fully-sampled w/ weighting



Retrospective
under-sampled
 $R=12$

Masked \rightarrow Abs \rightarrow Normalize

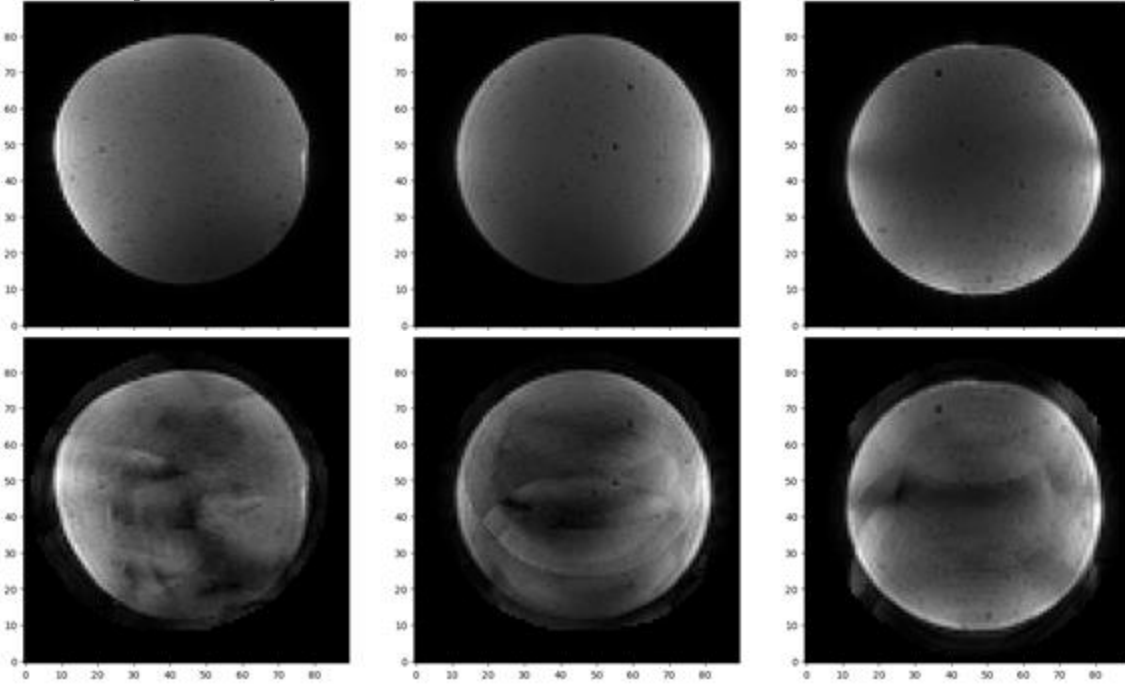
Sagittal View



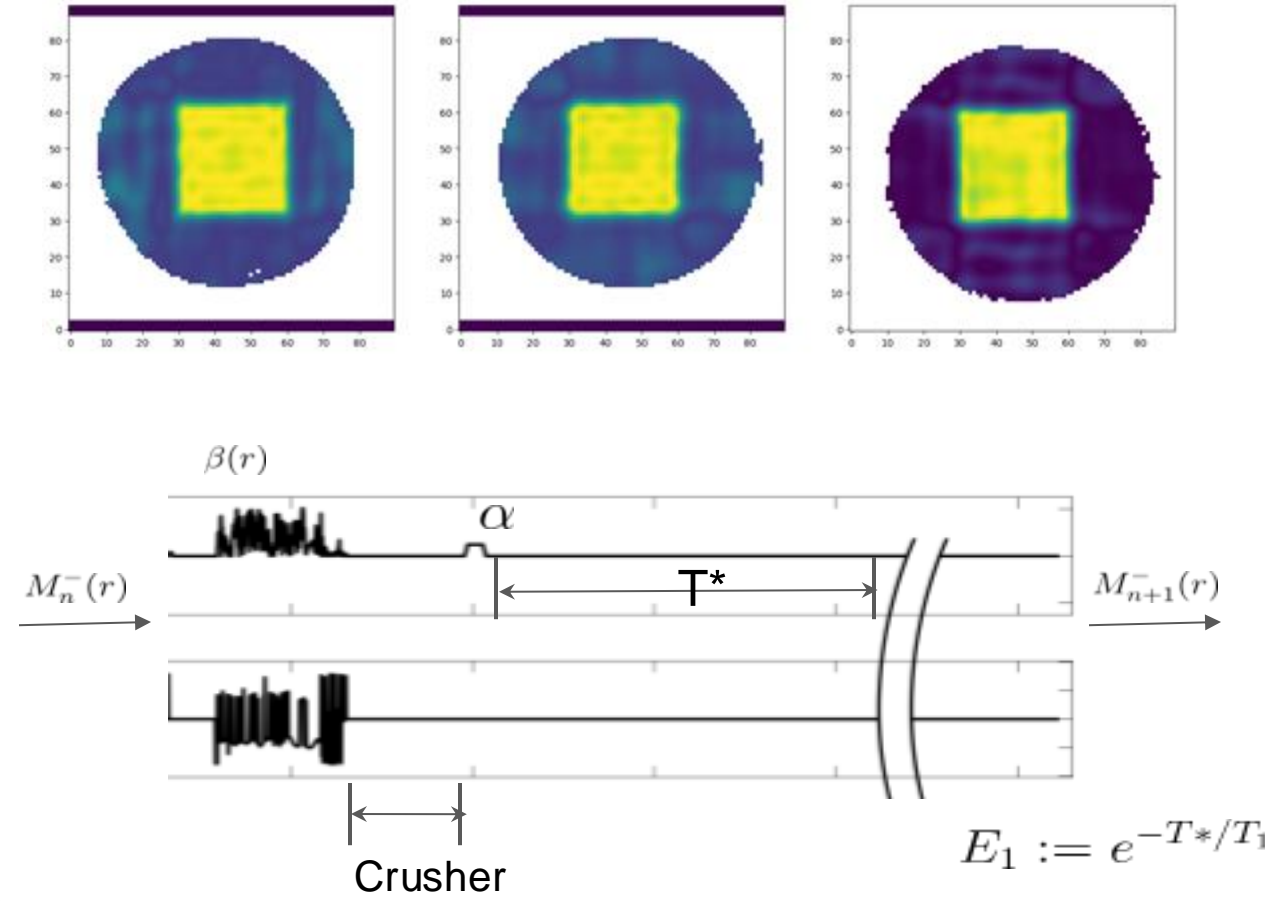
$$\lambda^* = \arg \min || \lambda v_1 - v_0 ||$$
$$= \frac{v_1^T v_0}{||v_0||_2^2}$$
$$\hat{v}_1 = v_1 \cdot \lambda^*$$

Data from another Scan

Fully-sampled



Prospective under-sampled R=12



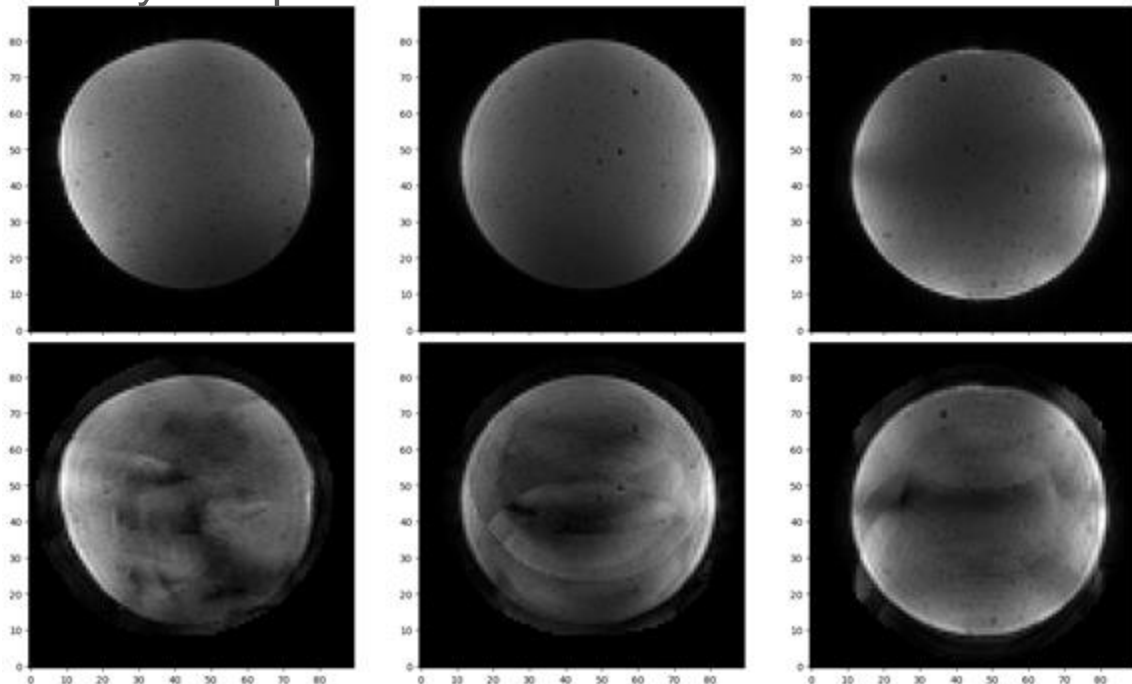
$$M_{SS} = \frac{M_0(1 - E_1)}{1 - \cos\beta(r)\cos\alpha E_1} \cdot \cos\beta(r)$$

$$M_{SS}^0 = \frac{M_0(1 - E_1)}{1 - \cos\alpha E_1}$$

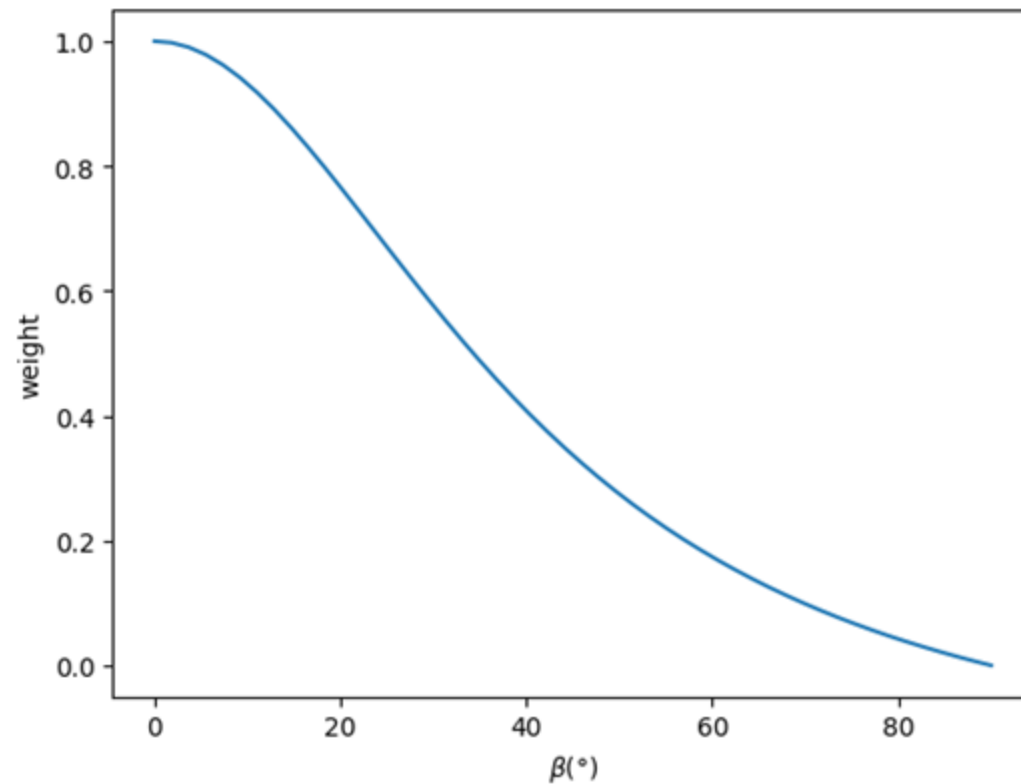
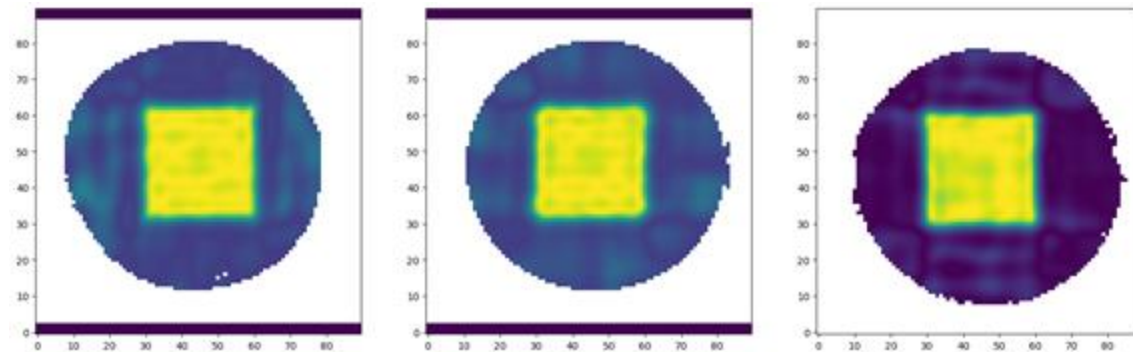
$$w := M_{SS}(r) / M_{SS}^0$$

Simulated tailored EXT

Fully-sampled

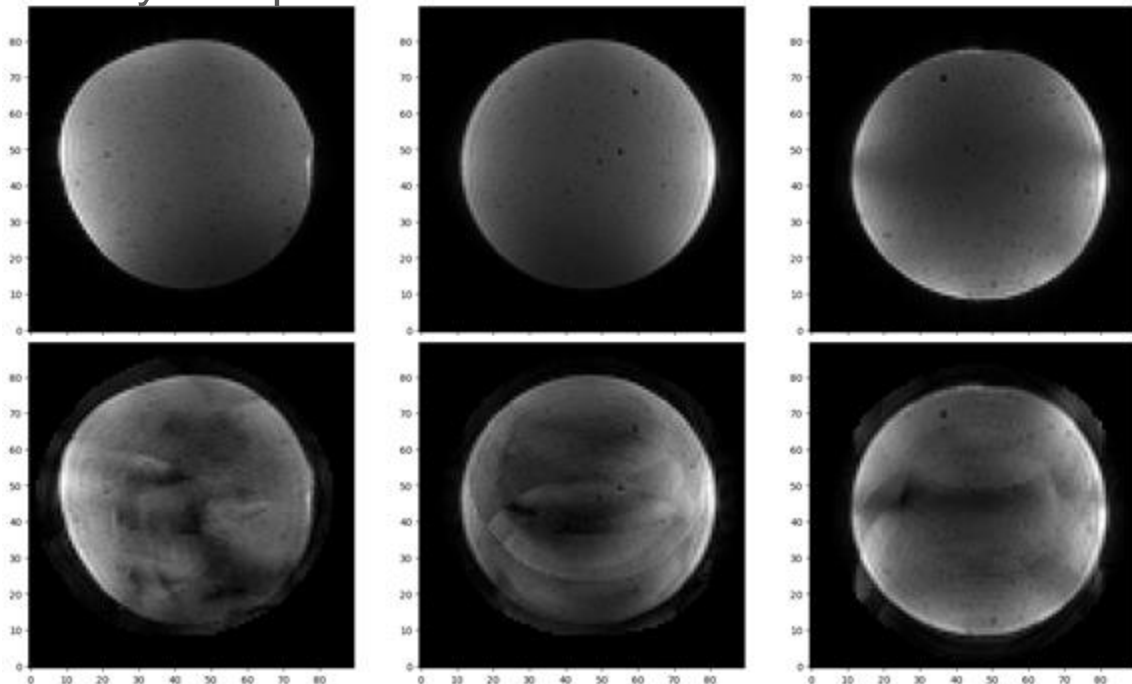


Prospective under-sampled R=12



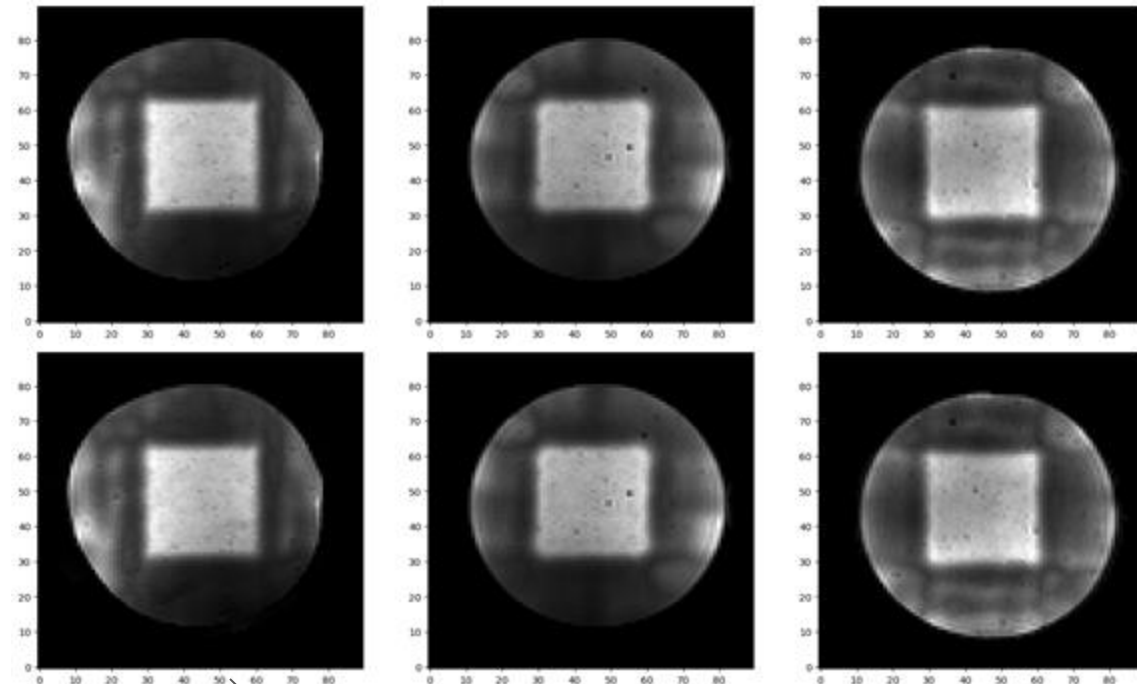
Simulated tailored EXT

Fully-sampled



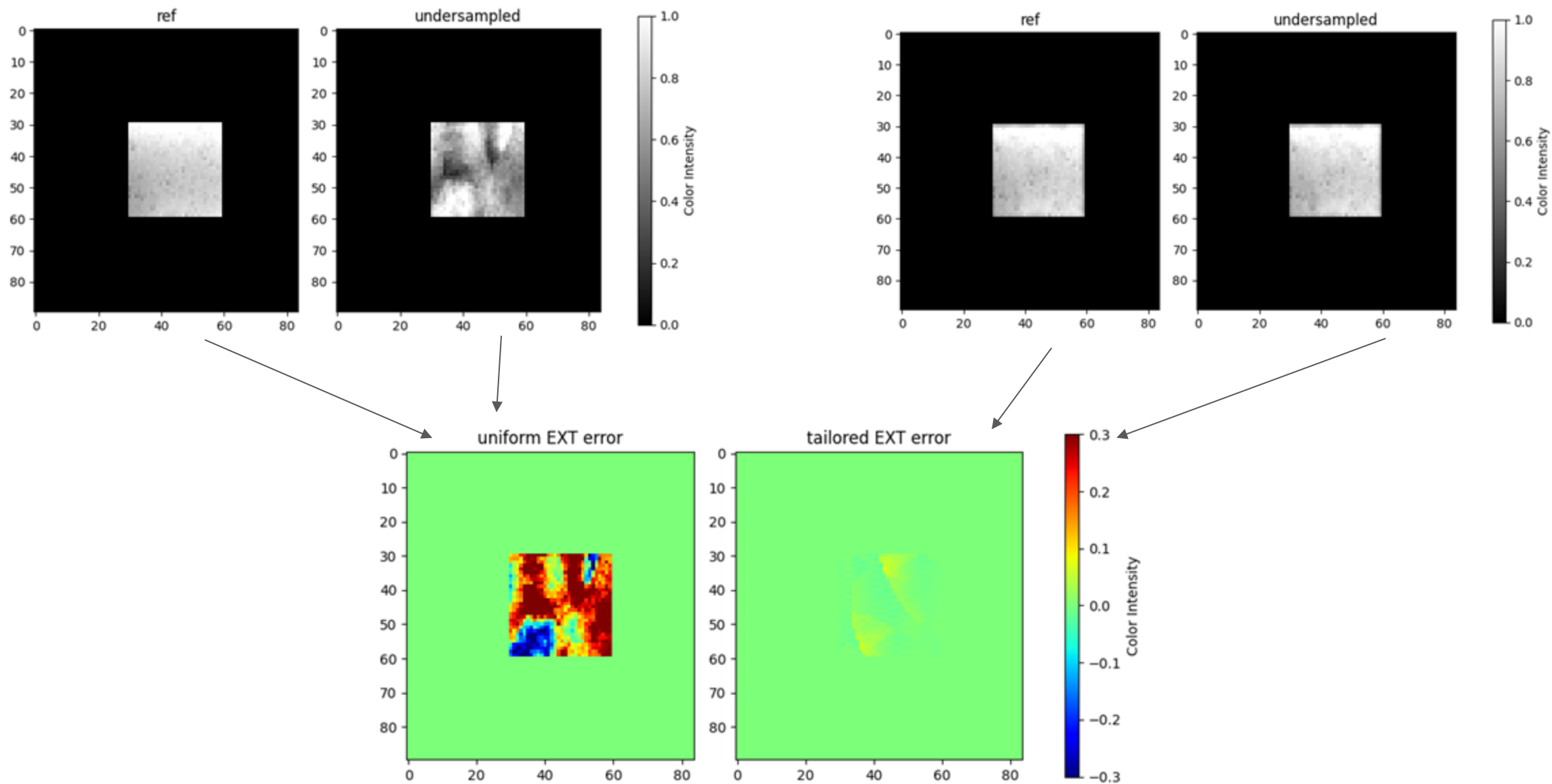
Prospective under-sampled R=12

Fully-sampled w/ simulated tEXT



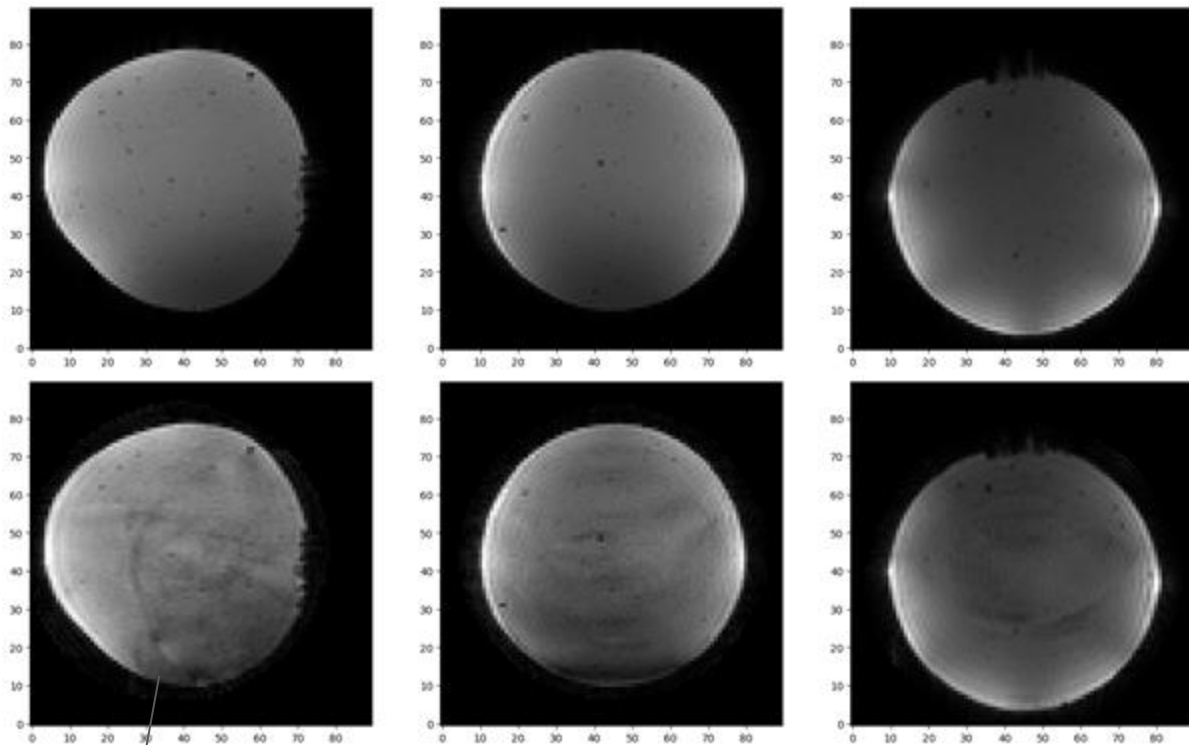
Retrospective under-sampled R=12

Simulated tailored excitation (tEXT)



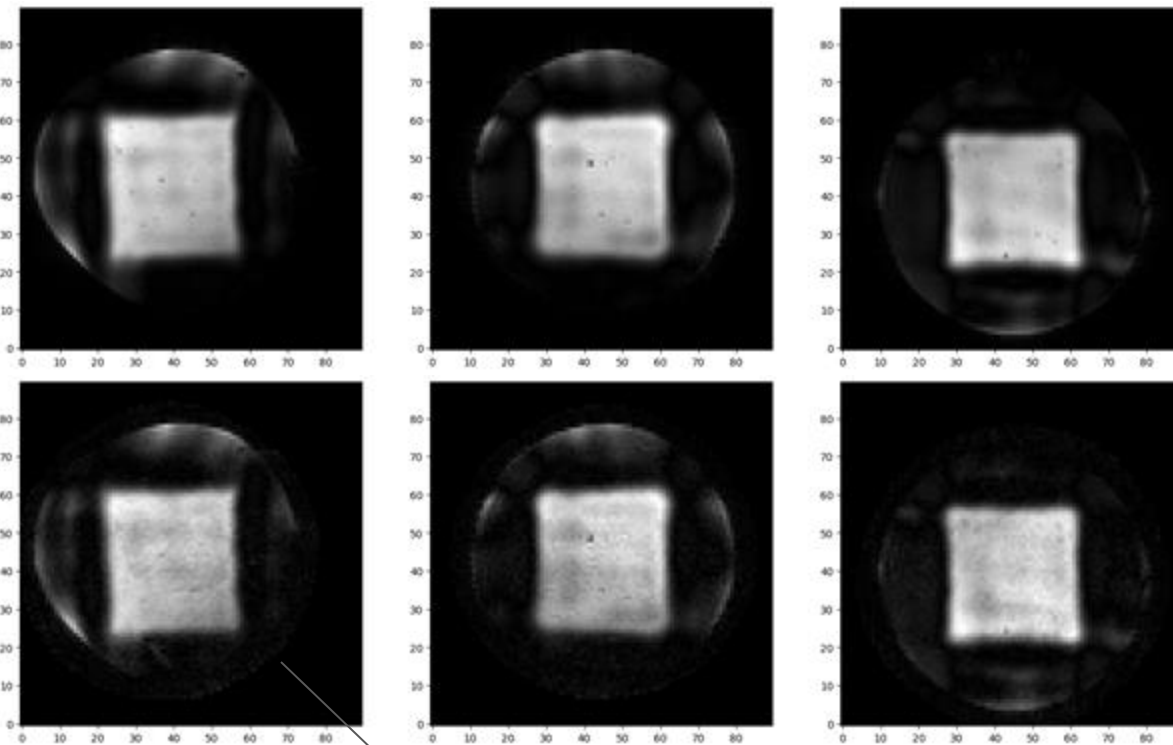
Real tailored excitation (tEXT)

Fully-sampled



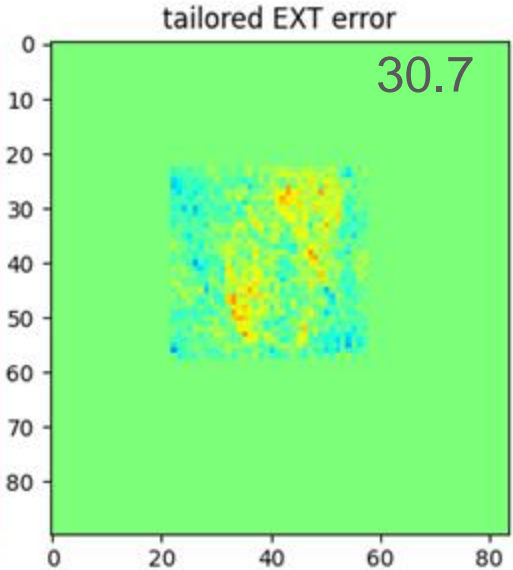
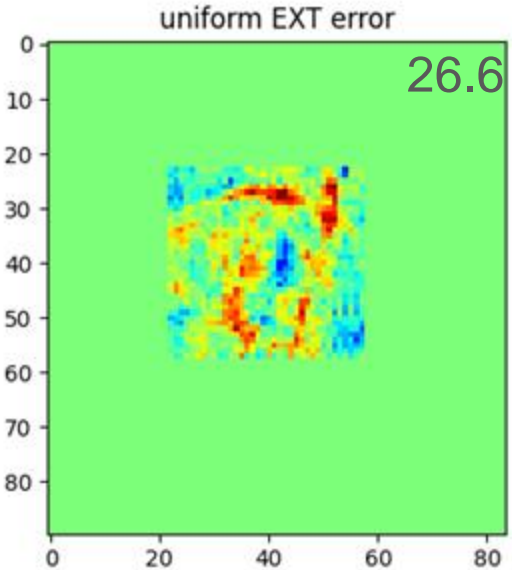
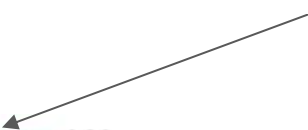
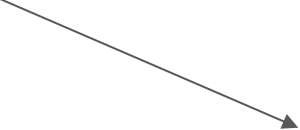
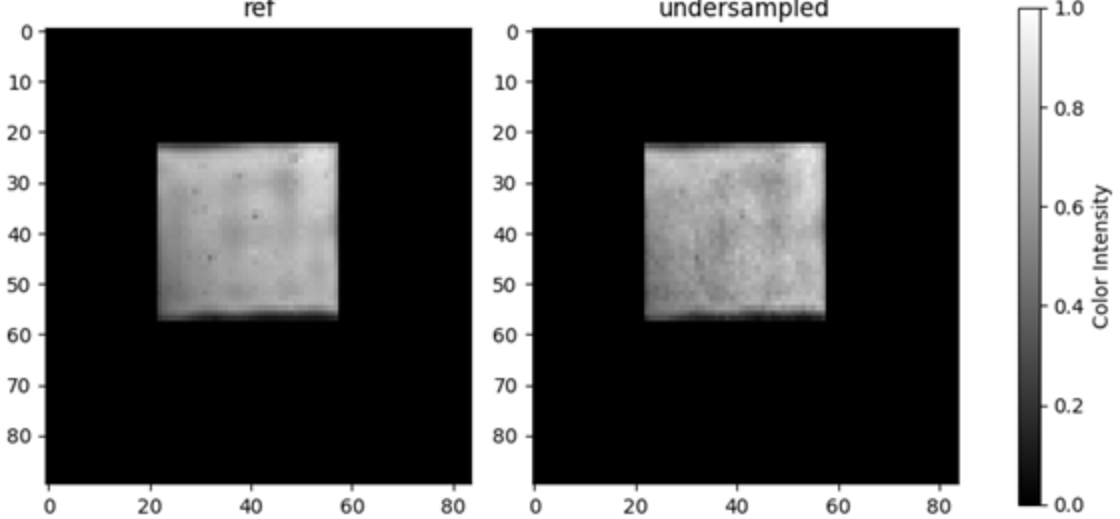
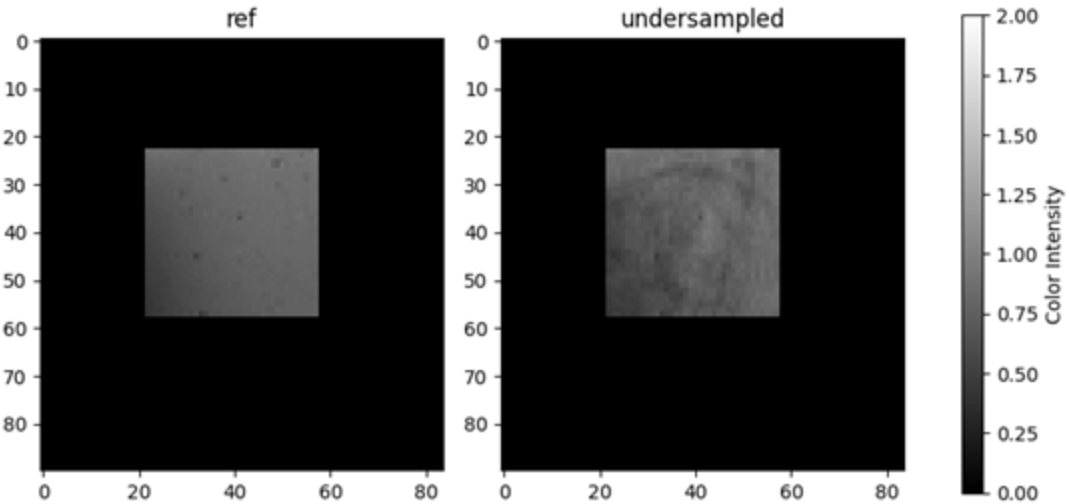
Prospective under-sampled R=12

Fully-sampled w/ (real) tEXT



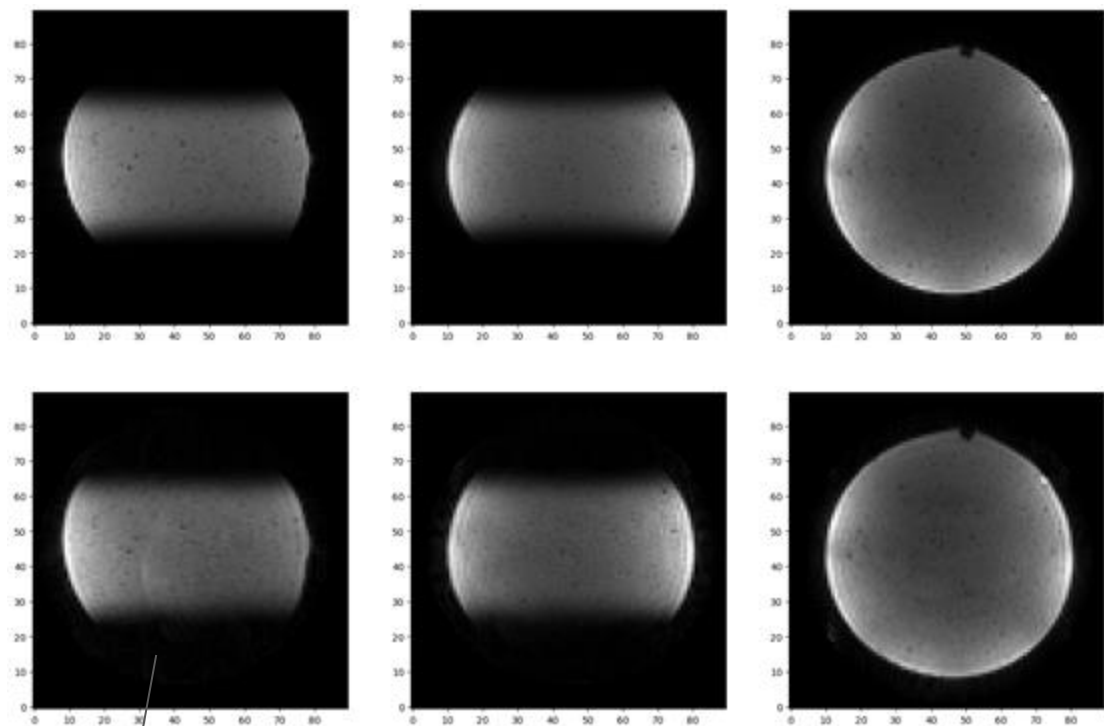
Prospective under-sampled R=12

Real tailored EXT



2D OVS + 1D EXT (20241122)

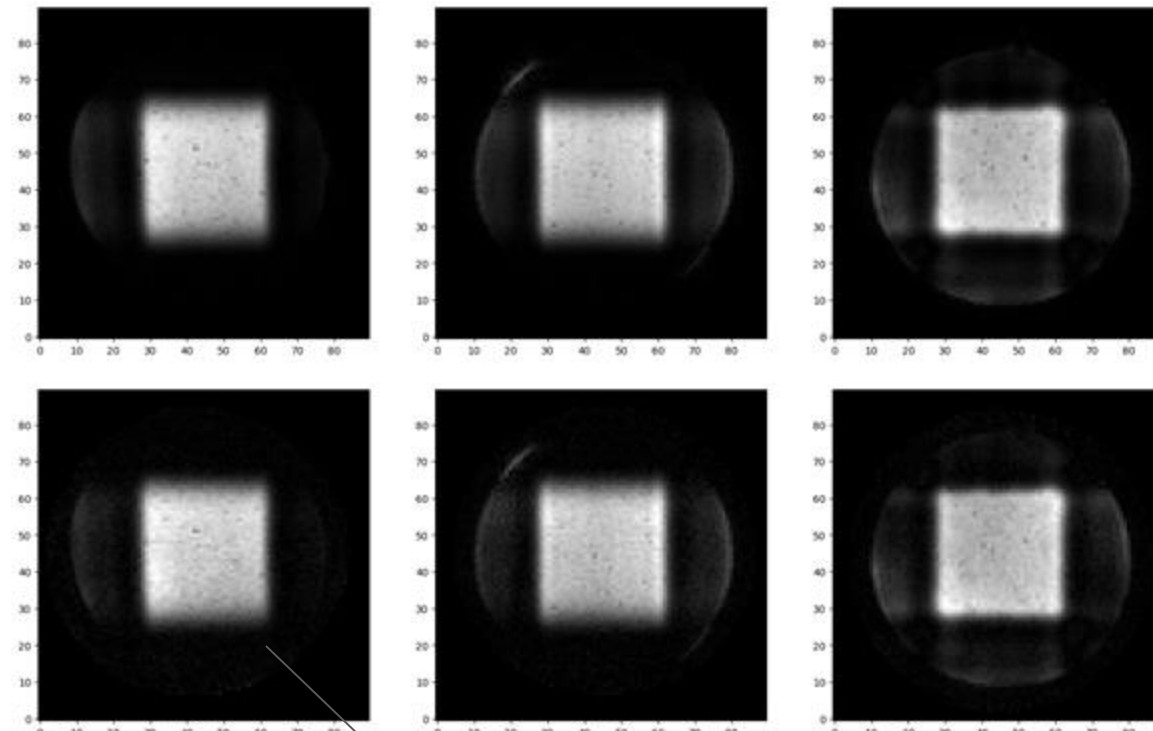
Fully-sampled



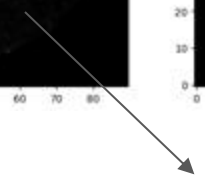
Prospective under-sampled R=12



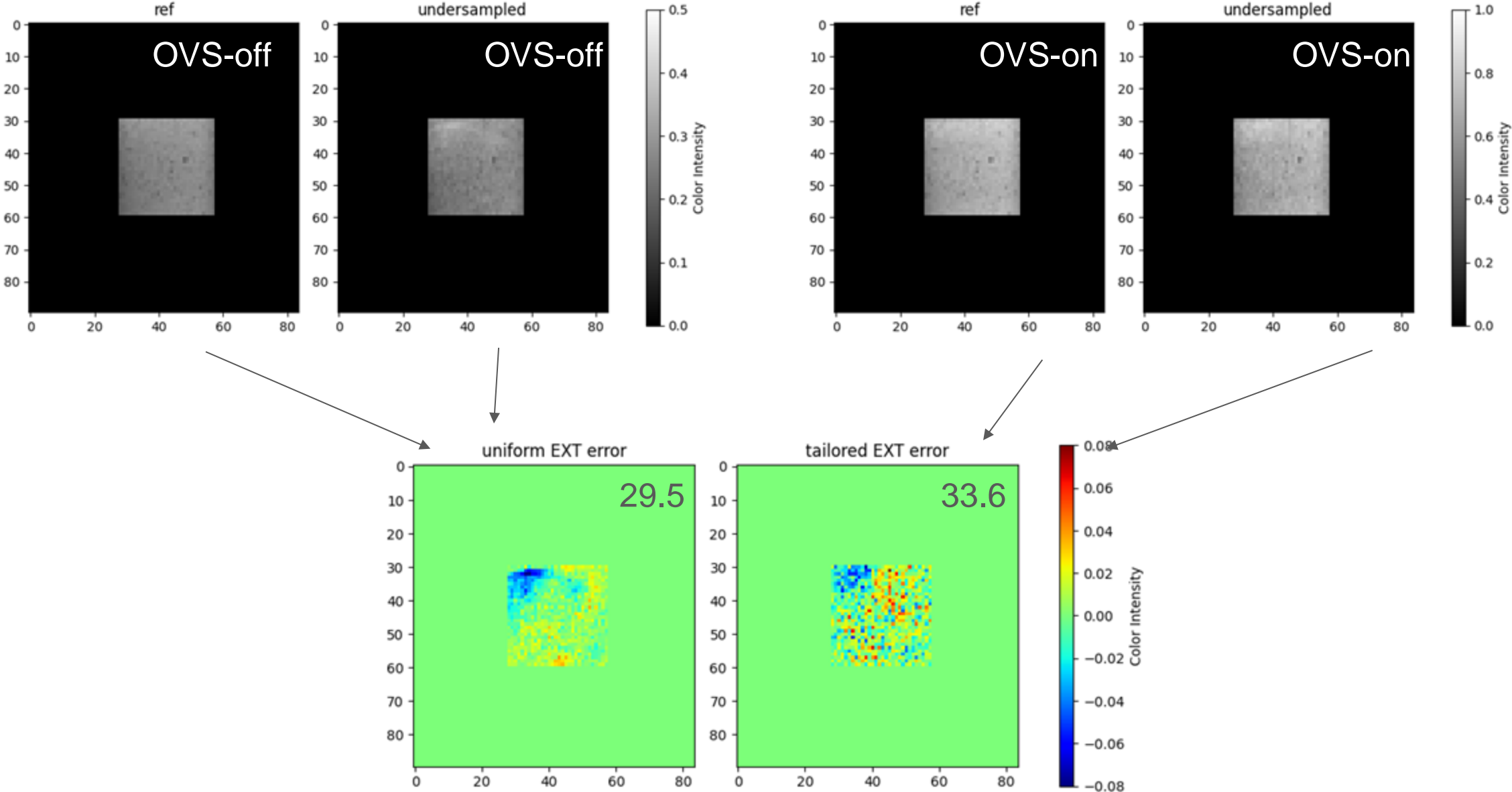
Fully-sampled w/ (real) tEXT



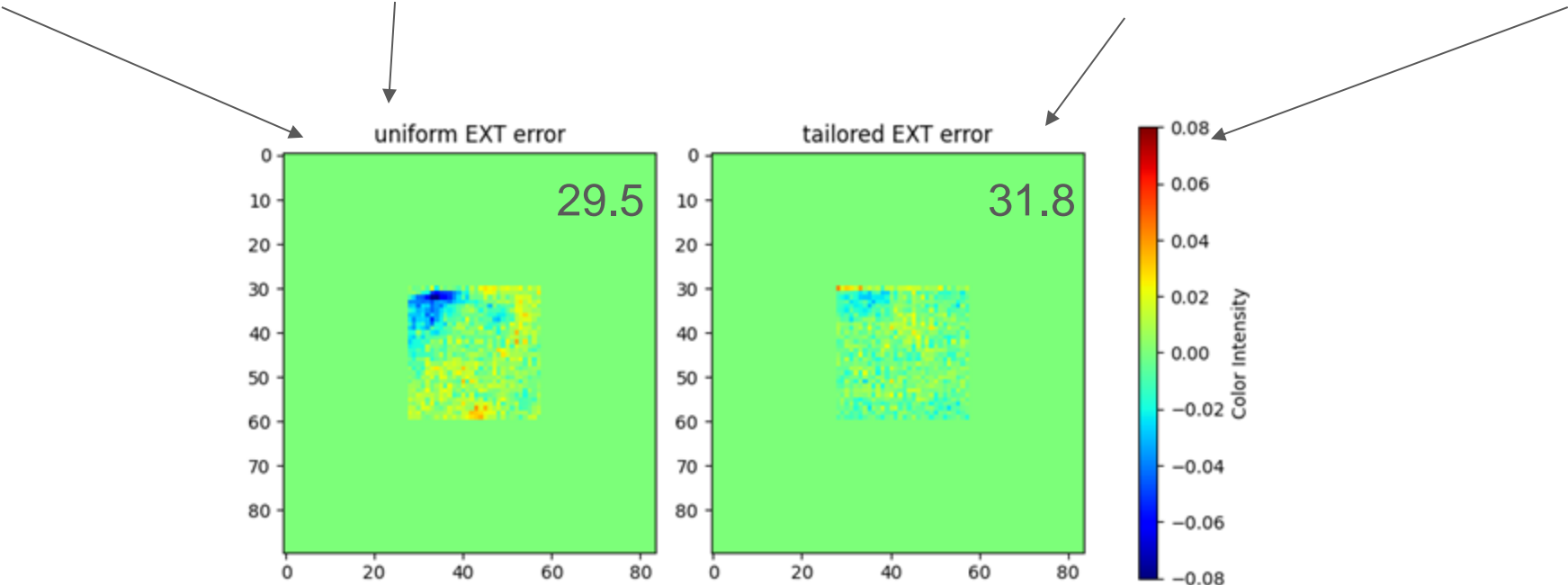
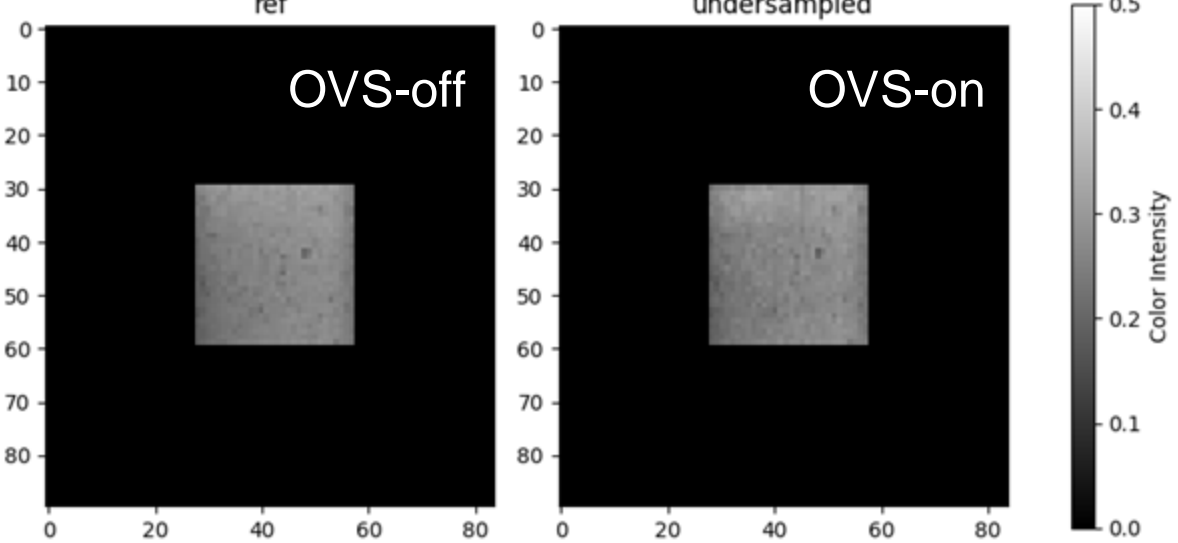
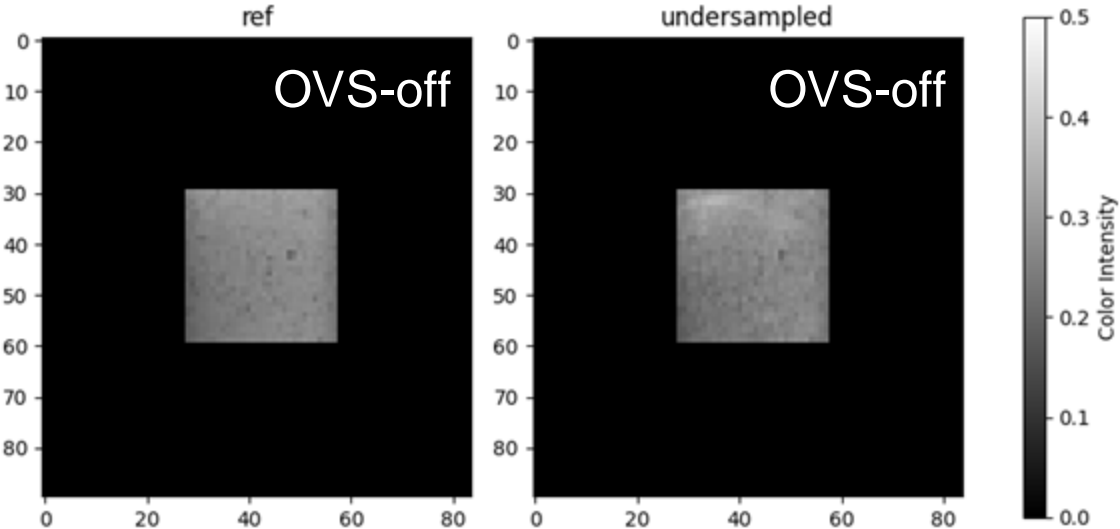
Prospective under-sampled R=12



2D OVS + 1D EXT (20241122)

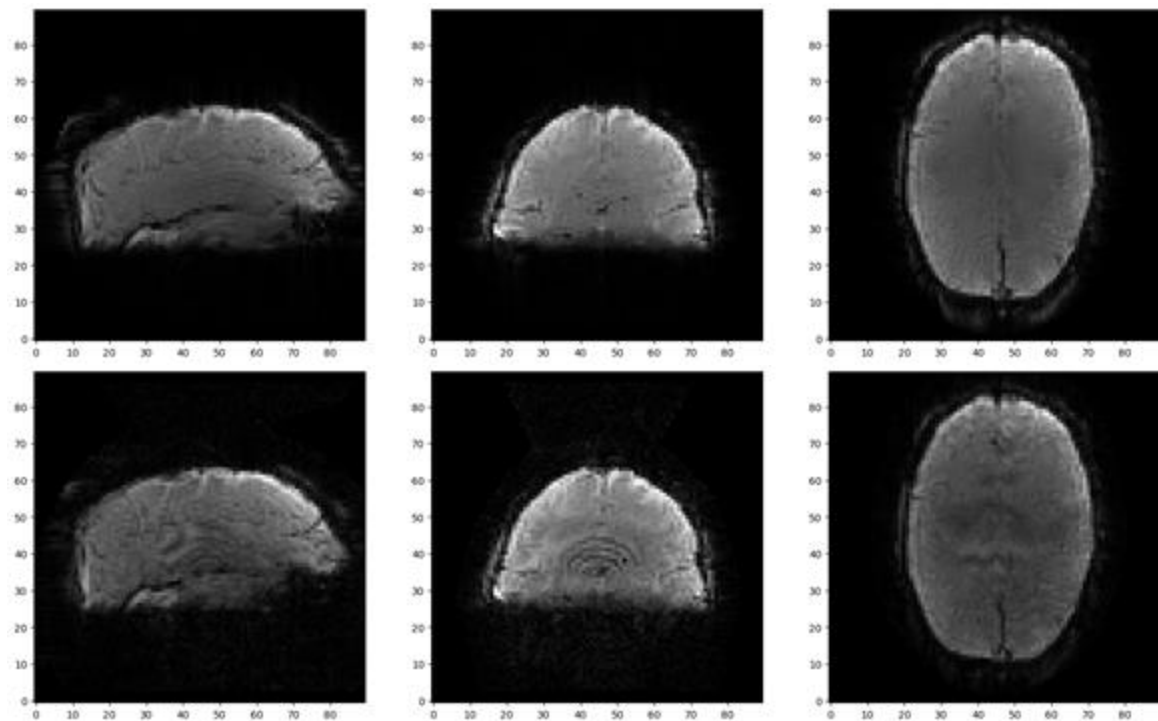


2D OVS + 1D EXT (20241122)



In Vivo (20241124)

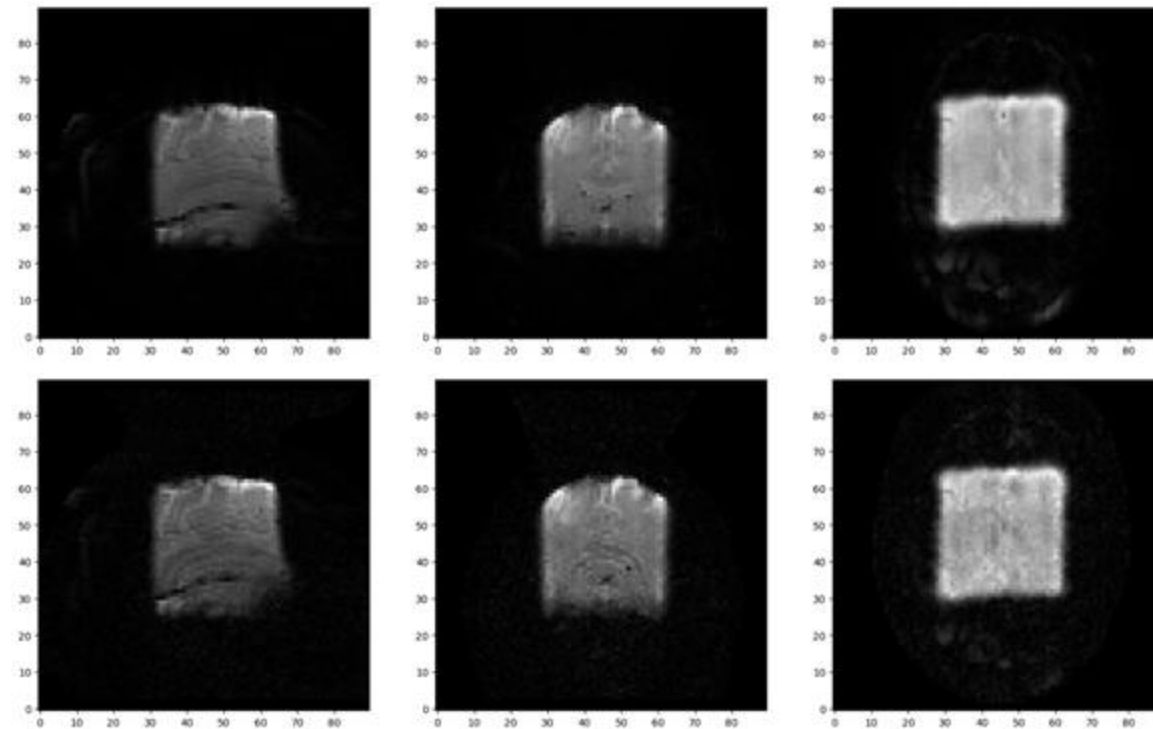
Fully-sampled



Prospective under-sampled R=12



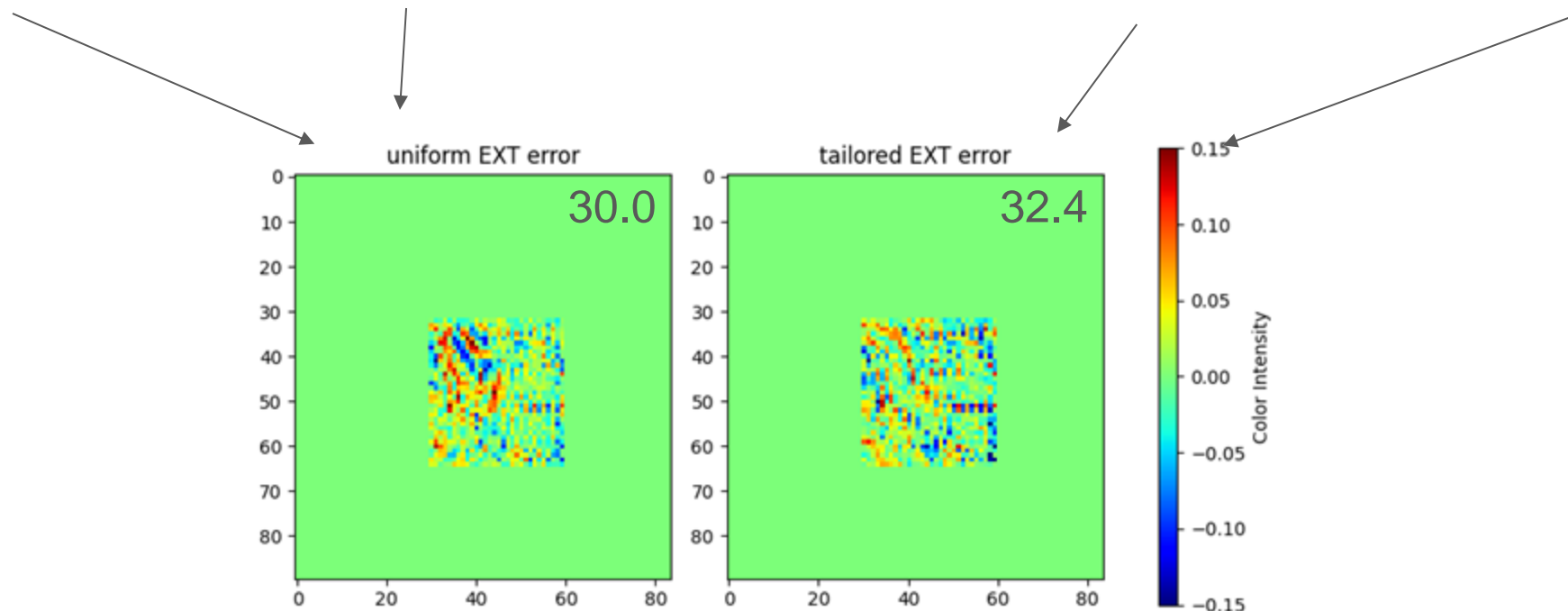
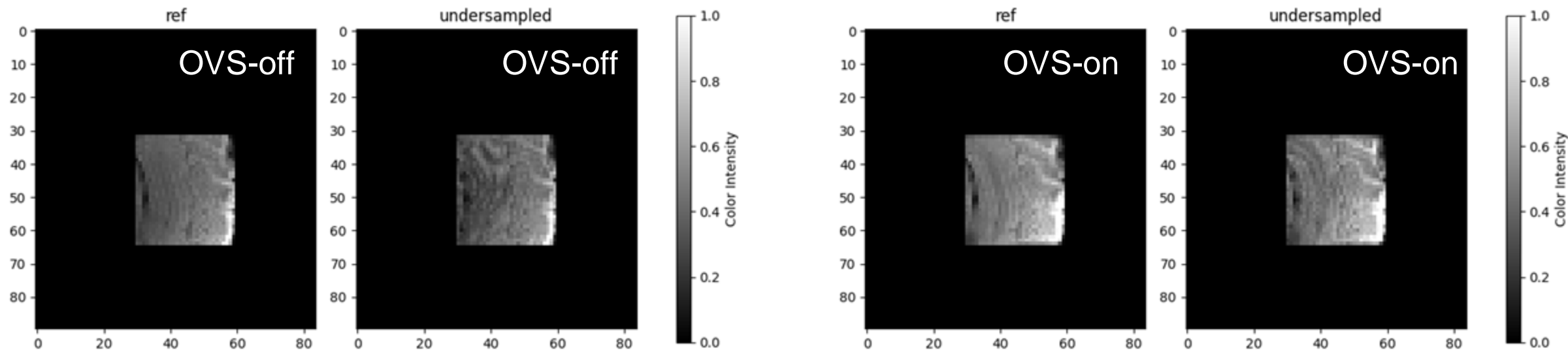
Fully-sampled w/ (real) tEXT



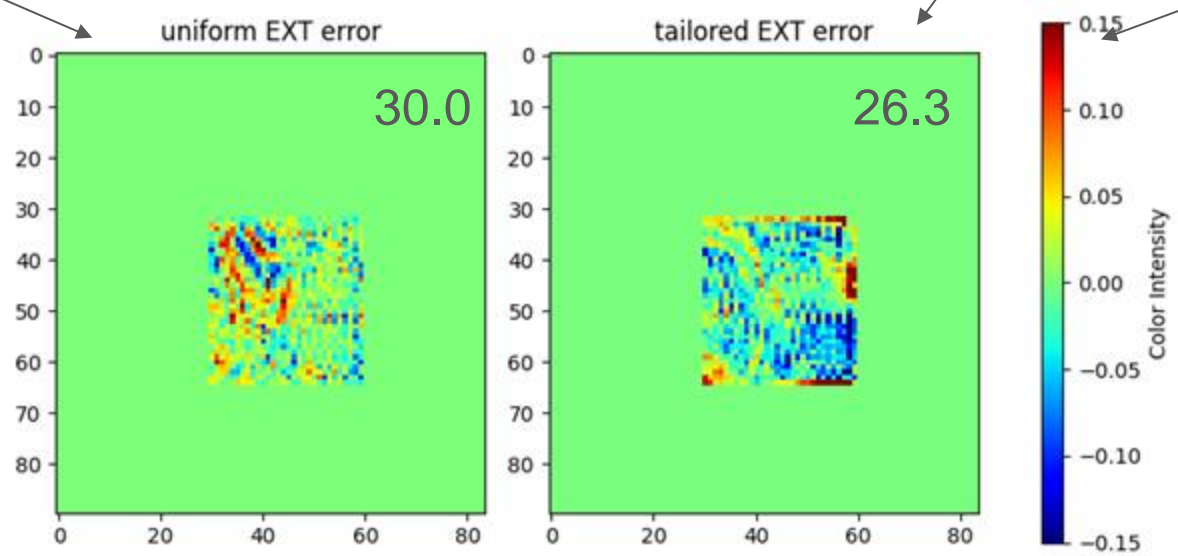
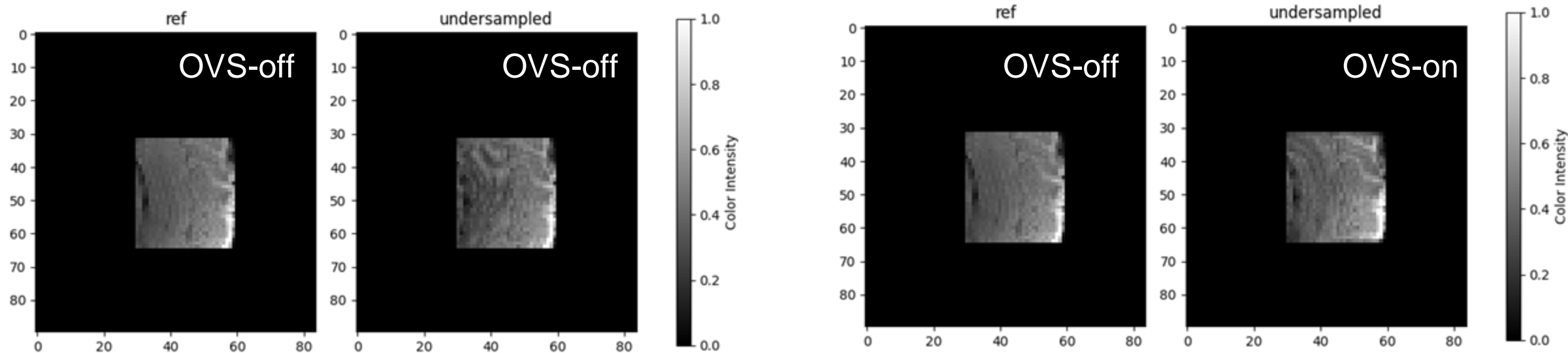
Prospective under-sampled R=12



In Vivo (20241124)



In Vivo (20241124)



IV excitation is not uniform for OVS-on

Next Step

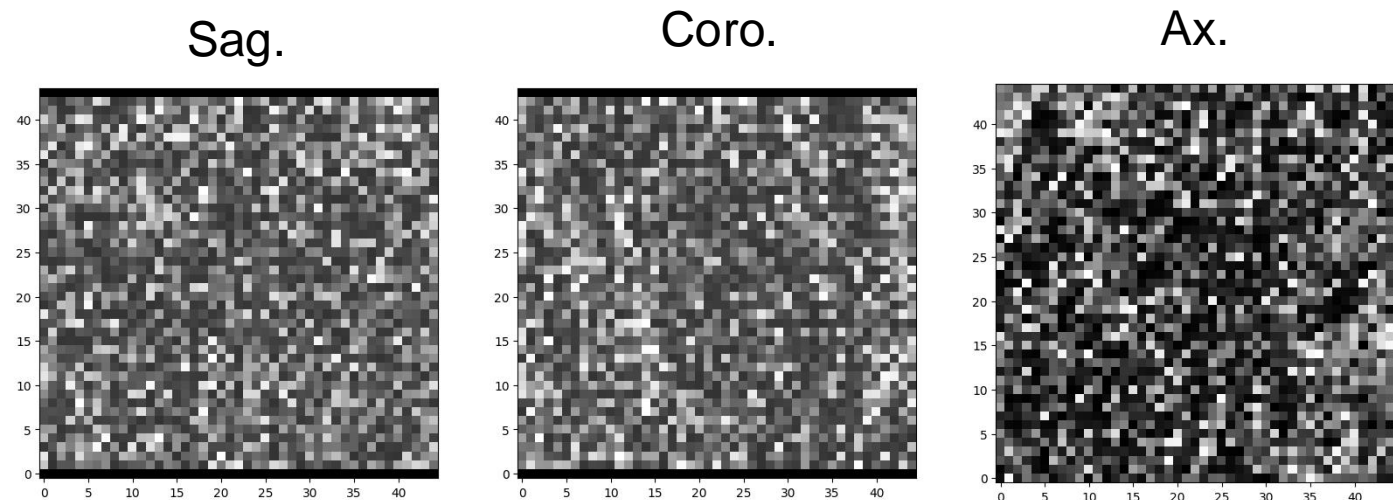
$$\mathcal{L} = \left\| I_{FS}(\text{OVSon}) - I_{US}(\text{OVSon}) \right\|_2$$

- Design tRF based on ROI image quality (reduce aliasing as goal)

Next Step

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- Design tRF based on ROI image quality (reduce aliasing as goal)
- One Caveat: If only use ROI error between fully-sampled and under-sampled OVS-on images as loss, the optimized excitation pattern might be totally random!



Next Step

$$\mathcal{L} = \left\| I_{FS}(\text{OVSON}) - I_{US}(\text{OVSON}) \right\|_2 + \lambda \cdot \mathcal{P}(|M_z^{tgt} - M_z^{tEXT}|_{ROI})$$

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- Enforce IV excitation uniformity as well, via, e.g., penalization

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- Enforce IV excitation uniformity as well, via, **e.g., penalization**
- Optimize sampling scheme in synergy with the tRF