## Closed-loop fMRI at the mesoscopic scale of columns and layers: Can we do it and why would we want to?

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## Supplementary appendix: Calculation of tSNR dependence on voxel size and number of measurements

We base our SNR calculations on the noise model of [1]. Voxel independent physiological  $\sigma_p$  and voxel volume dependent thermal noise  $\sigma_0$  are given by  $\sigma_p = \lambda$  and  $\sigma_0 = 1/\kappa V$ , with total time-course SNR  $tSNR = \frac{1}{\sqrt{\sigma_0^2 + \sigma_p^2}} = \frac{\kappa V}{\sqrt{1 + \lambda^2 \kappa^2 V^2}}$ . Triantafyllou et al. [1],

measured SNR as a function of voxel volume for different magnetic field strengths which can be used to estimate  $\lambda$  and  $\kappa$ . Chaimow et al. [2] derived how to translate  $\kappa$  for different TRs using grey matter  $T_1$  values. Assuming uncorrelated thermal noise,

averaging of  $N_t$  consecutive measurements in time and  $N_v$  voxels in space results in a

reduced thermal noise component  $\sigma_0^{N_t,N} = \frac{\sigma_0}{\sqrt{N_v N_t}} = \frac{1}{\kappa V \sqrt{N_v N_t}}$ 

Furthermore, we assume physiological noise to be approximately homogeneous in space (at small spatial scales) but autocorrelated in time with autocorrelation

$$E\left[\epsilon_{t_1}^{p}\epsilon_{t_2}^{p}\right] = \frac{\sigma_p^2 \exp(-TR|t_1 - t_2|)}{15} [3],$$

where  $\epsilon_t^p$  is the noise component of measurement number *t*. Assuming zero mean noise, we obtain a reduced physiological noise level after averaging  $N_t$  measurements

according to

$$\left(\sigma_{p}^{N_{t}}\right)^{2} = E\left[\left(\frac{1}{N_{t}}\sum_{t=1}^{N_{t}}\epsilon_{t}^{p}\right)^{2}\right] = \frac{1}{N_{t}^{2}}\left(\sum_{t_{1}=1}^{N_{t}}\sum_{t_{2}=1}^{N_{t}}E\left[\epsilon_{t_{1}}^{p}\epsilon_{t_{2}}^{p}\right]\right) = \frac{1}{N_{t}^{2}}\sum_{t_{1}=1}^{N_{t}}\sum_{t_{2}=1}^{N_{t}}\frac{\sigma_{p}^{2}\exp(-TR|t_{1}-t_{2}|)}{15}.$$

Taken together we can calculate total noise as follows. We start with fitted values of  $\kappa$  and  $\lambda$  that were estimated for different magnetic field strength and a TR of 5.4 s [1,4] and modify  $\kappa$  to reflect a TR of 2 s, which is more typical of a current fMRI study [2].

Combining independent thermal and physiological noise components we obtain

$$SNR^{N_{t'}N} = \frac{1}{\sqrt{\left(\sigma_{p}^{N_{t}}\right)^{2} + \left(\sigma_{0}^{N_{t'}N}\right)^{2}}} = \frac{\kappa(TR)V_{\sqrt{N_{v}N_{t}}}}{\sqrt{1 + \kappa^{2}(TR)V^{2}N_{v}N_{t}}\lambda^{2}\left(\frac{1}{N_{t}^{2}}\sum_{t_{1}=1}^{N_{t}}\sum_{t_{2}=1}^{N_{t}}\frac{\exp(-TR|t_{1}-t_{2}|)}{15}\right)}}$$

For 3T we used:  $\kappa = 6.6567$ ,  $\lambda = 0.0129$ ,  $T_1 = 1.607$ ; and for 7T:  $\kappa = 9.9632$ ,  $\lambda = 0.0113$ ,  $T_1 = 1.939$  [4].

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3.Purdon PL, Weisskoff RM. 1998 Effect of temporal autocorrelation due to physiological noise and stimulus paradigm on voxel-level false-positive rates in fMRI. *Human Brain Mapping* **6**, 239–249.

4.Chaimow D, Uğurbil K, Shmuel A. 2018 Optimization of functional MRI for detection, decoding and high-resolution imaging of the response patterns of cortical columns. *NeuroImage* **164**, 67–99. (doi:10.1016/j.neuroimage.2017.04.011)