# Inference of Latent Neural Field Intensities from Spatiotemporal Point-Process Observations

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## 1. 3-State model for waves in excitable medium

Three-state neural field model, Buice & Cowan '09 (1): complex wave patterns without inhibitory cells.

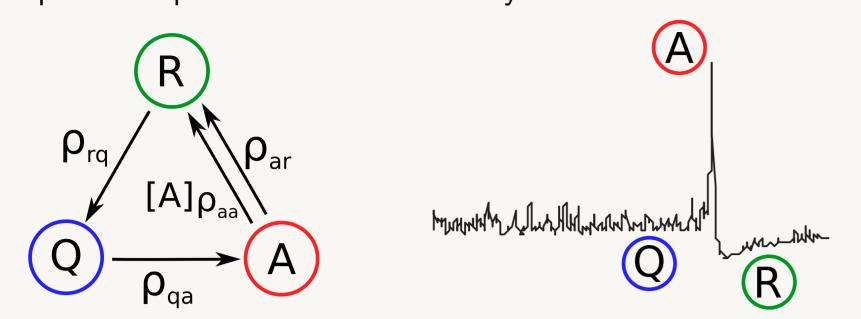
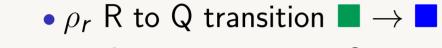


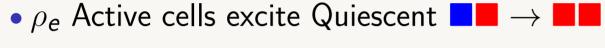
Figure 1. Quiescent-Active-Refractory (QAR) model of neural waves. The Q-A-R states (1) correspond to the Critical-Active-Stable retinal wave model of Hennig et al. '09 (2) and the Susceptible-Infected-Recovered model in epidemiology.

#### 3 states

- Q Quiescent
- A Active
- R Refractory



•  $\rho_a$  A to R transition  $\blacksquare \rightarrow \blacksquare$ 



•  $\rho_q$  Spontaneous spiking; Q to A  $\longrightarrow$ 

**Mean-field dynamics** (exclude spontaneous  $Q \rightarrow A$ )

4 rate parameters

$$\dot{Q} = -\rho_e A Q + \rho_r R$$
 $\dot{A} = -\rho_a A + \rho_e A Q$ 
 $\dot{R} = -\rho_r R + \rho_a A$ 

Spontaneous  $Q \rightarrow A$  sampled as shot noise (Poisson).

#### 2. Spatial system

Let fields depend on coordinates (x, y) and define a lateral excitation kernel k with radius  $\sigma_i$  (Nonlocal interactions)

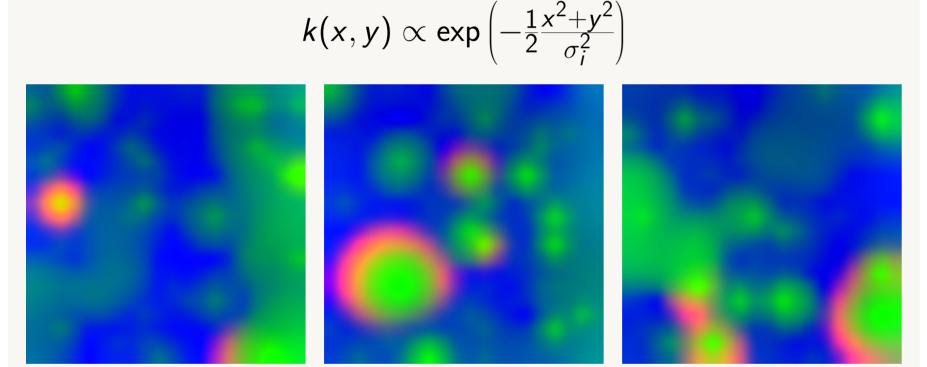


Figure 2. 3-state model can exhibit self-organized wave phenomena. Simulated on a  $[0,1]^2$  unit interval using 20x20 grid,  $\sigma$ =0.04,  $\rho_a$ =0.1,  $\rho_r$ =10<sup>-3</sup>,  $\rho_e$ =0.4. Spontaneous excitation rate  $\rho_q$ =0.05. A finite threshold of 10<sup>-3</sup> avoids widespread spontaneous excitation. Colors: Quiescent Active Refractory

#### 3. Recover latent fields from spikes: Bayesian filtering

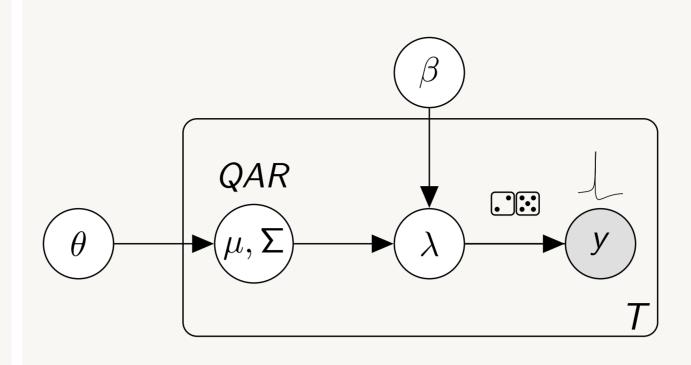


Figure 3. Hidden Markov model for latent neural **fields.** For all time-points T, state transition parameters  $\theta = (\rho_q, \rho_a, \rho_r, \rho_e, \sigma)$  dictate the evolution of a multivariate Gaussian model  $\mu$ ,  $\Sigma$  of latent fields Q, A, R. Observation model  $\beta$  is a linear map with adjustable gain and threshold, and reflects how field A couples to firing intensity  $\lambda$ . Pointprocess observations (spikes) y are Poisson with intensity  $\lambda$ .

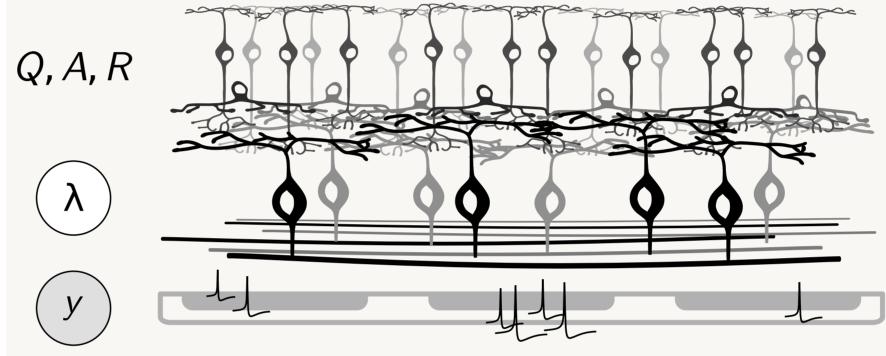
### **Predict state:**

- Multivariate Gaussian state-space model  $\mu = (Q, A, R)$ , covariance  $\Sigma$
- ullet Integrate forward  $\mu$  mean-field equations
- ullet Covariance  $\Sigma$  evolves according to the system Jacobian J
- Similar to continuous-time extended Kalman filter  $\dot{\Sigma} = J\Sigma + \Sigma J^T + \Sigma_{\text{noise}}$

#### Measurement:

- Refine estimate using spiking observations
- Spikes: Poisson events with intensity  $\lambda = mA + b$
- Posterior is proportional to product of predicted state and data likelihood
- Laplace approximation (gradient descent; constrain to positive field intensities)

# Test case: developmental retinal waves



Bipolar and amacrine cells (generate waves)

**Retinal Ganglion Cells** 

Multi-electrode array (spiking observations)

Figure 4. Illustration of inner retina and recording setup. Spontaneous retinal waves are generated in a layer of laterally interconnected amacrine cells. These waves activate Retinal Ganglion Cells (RGCs), the output cells of the retina. RGC electrical activity is recorded via a  $64\times64$  multi-electrode array with 50 µm spacing.

## **High-density multielectrode array** recordings of retinal waves

- 4096-electrode arrays, 42 μm spacing (3)
- Recordings courtesy of the Sernagor lab (4, 5)
- Spontaneous waves during development (6)
- Small events divide retina into refractory patches
- Rare large events sweep across the retina
- Self-organized structure at multiple scales (2)

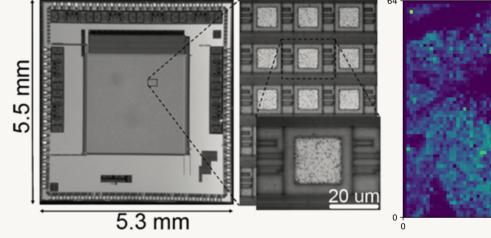
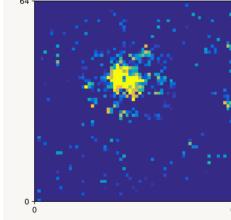
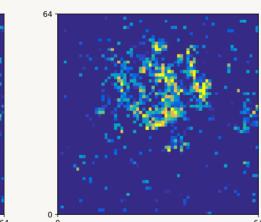
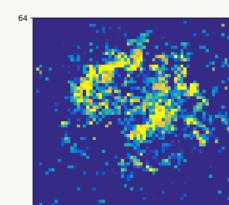
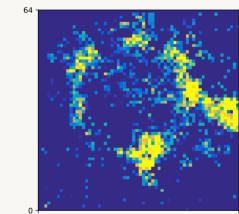


Figure 5. 4096-electrode array. Left: Array (3). Right: Spikes recorded in a single session.









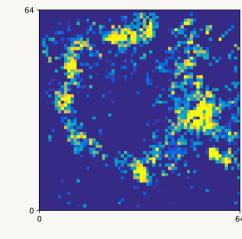


Figure 6. Example wave event, spike histograms in 1 s intervals.  $2.6 \times 2.6$  cm<sup>2</sup>. Mouse retina, postnatal day 11.

#### 5. In practice

## Numerically challenging:

- 3 states,  $10 \times 10$  grid  $\rightarrow$  300-D covariance matrix (4.5k entries)
- Avoid inverses: work with inverse covariance (precision) matrix
- Improve stability: Cholesky factorization, triangular system solvers
- Regularize state variance

# Performance e.g.:

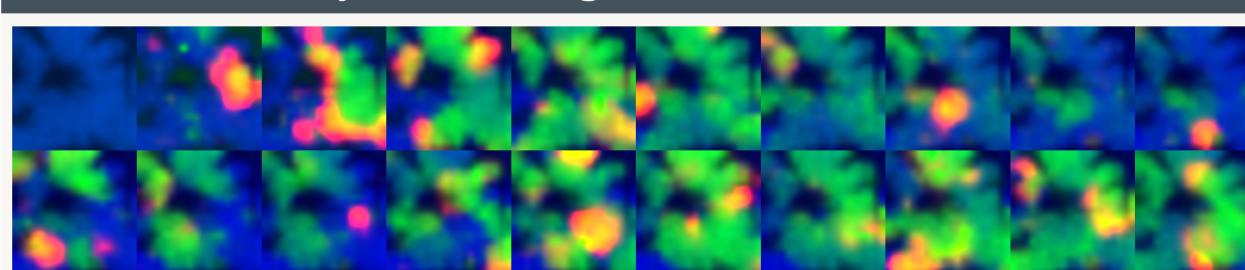
- 37 s to filter 25 minutes of retinal data,  $\Delta t=1$  s,  $\sim$ 40 samples/s
- 10×10 grid; Matlab implementation, 2.9 GHz 8-core Xeon CPU
- Complexity dominated by matrix multiplication

## **Fluctuations:**

- A model of fluctuations is needed to model uncertainty in state estimation
- Use a linear noise approximation of the original discrete system

$$\Sigma_{\text{noise}} = \begin{bmatrix} \rho_e A Q + \rho_r R & -\rho_e A Q & -\rho_r R \\ -\rho_e A Q & \rho_i A + \rho_e A Q & -\rho_i A \\ -\rho_r R & -\rho_i A & \rho_r R + \rho_i A \end{bmatrix}$$

#### 6. Bayesian filtering recovers latent states



**Figure 7. Filtering recovers wave states.** Brightness  $\propto$  cell density. Colors: Quiescent Active Refractory. Frames shown every 48 seconds;  $2.6 \times 2.6$  cm<sup>2</sup> area; postnatal day 10; Multiscale "forest-fire like" waves.

## Main points

- Spatiotemporal neural phenomena are complex: excitability, nonlinearity, refractoriness
- Previous spatiotemporal point-process inference procedures unsuitable (simple, linear)
- Three-state neural field model is suitable for inference
- Bayesian filtering recovers latent states, correlation structure, and model likelihood
- Future: model validation, forecasting, and control.





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