

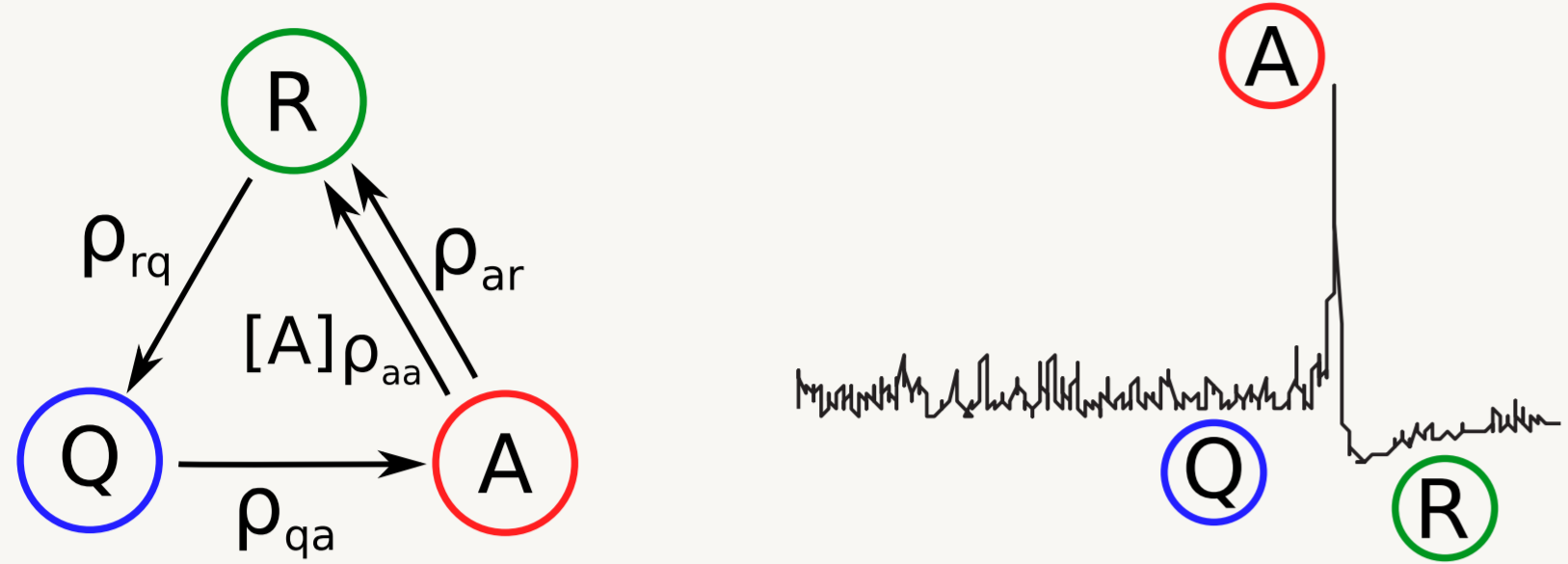
# Inference of Latent Neural Field Intensities from Spatiotemporal Point-Process Observations

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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

### 4 rate parameters

- $\rho_q$  Spontaneous spiking; Q to A ■  $\rightarrow$  ■
- $\rho_a$  A to R transition ■  $\rightarrow$  ■
- $\rho_r$  R to Q transition ■  $\rightarrow$  ■
- $\rho_e$  Active cells excite Quiescent ■ ■  $\rightarrow$  ■

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

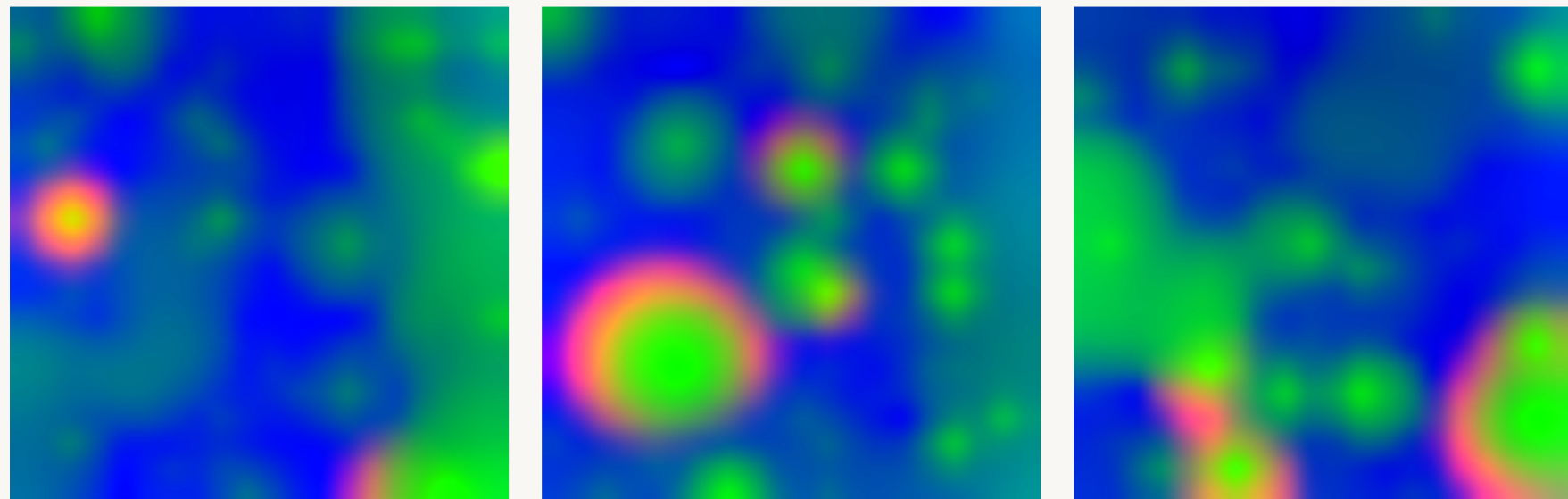
$$\begin{aligned}\dot{Q} &= -\rho_e A Q + \rho_r R \\ \dot{A} &= -\rho_a A + \rho_e A Q \\ \dot{R} &= -\rho_r R + \rho_a A\end{aligned}$$

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)

$$k(x, y) \propto \exp\left(-\frac{1}{2} \frac{x^2 + y^2}{\sigma_i^2}\right)$$



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

## 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.:

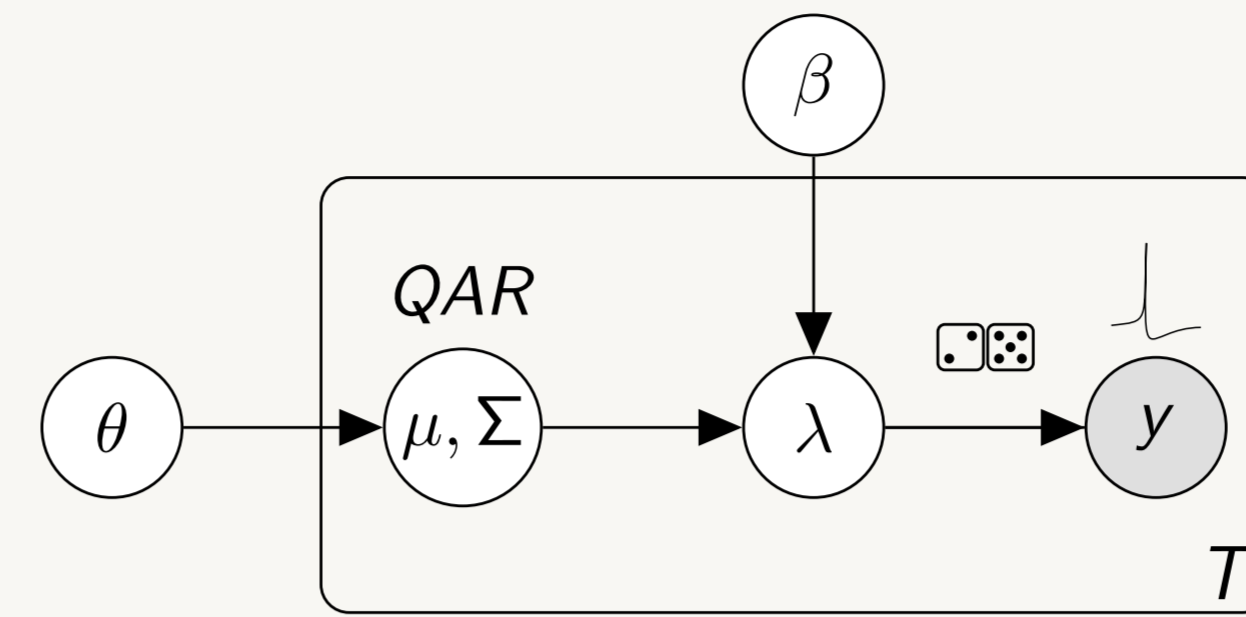
- 37 s to filter 25 minutes of retinal data,  $\Delta t = 1$  s,  $\sim 40$  samples/s
- $10 \times 10$  grid; Matlab implementation, 2.9 GHz 8-core Xeon CPU
- Complexity dominated by matrix multiplication

### Fluctuations:

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

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

## 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  $\lambda$ . Point-process observations (spikes)  $y$  are Poisson with intensity  $\lambda$ .

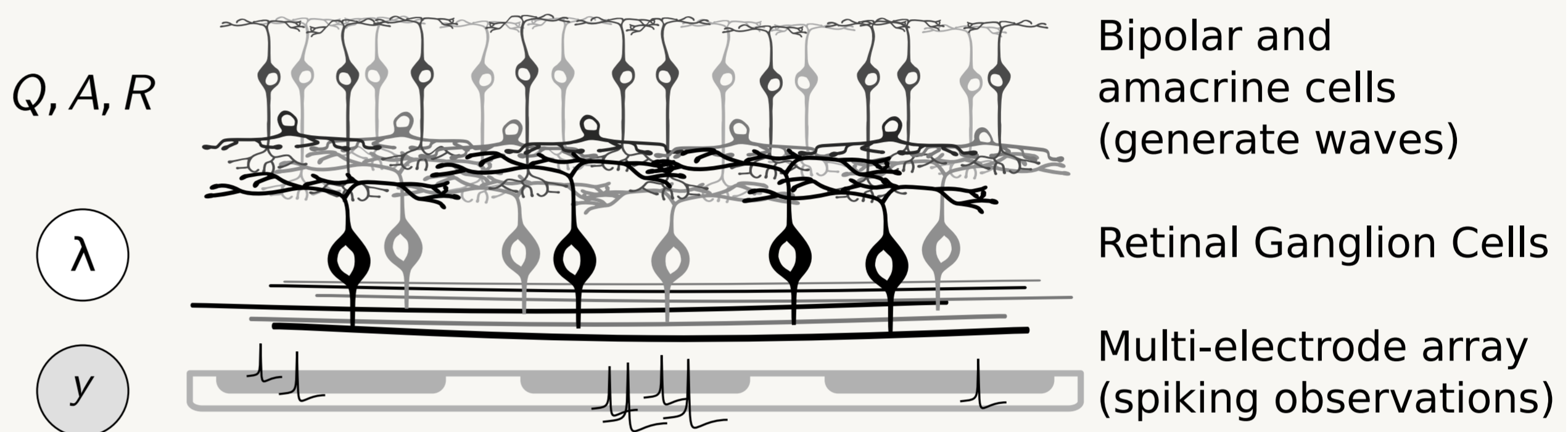
### Predict state:

- Multivariate Gaussian state-space model  $\mu = (Q, A, R)$ , covariance  $\Sigma$
- Integrate forward  $\mu$  mean-field equations
- 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)

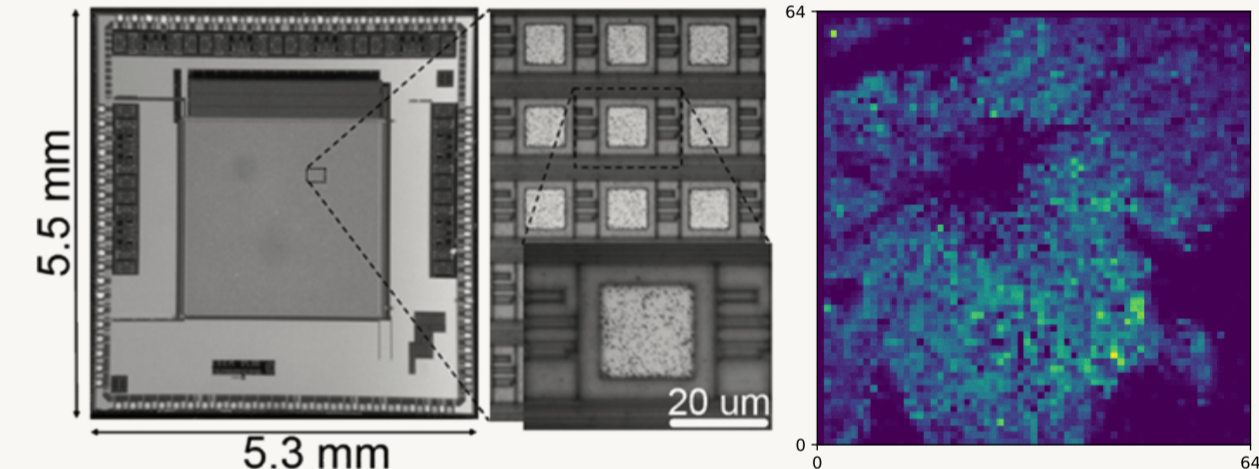
## 4. Test case: developmental retinal waves



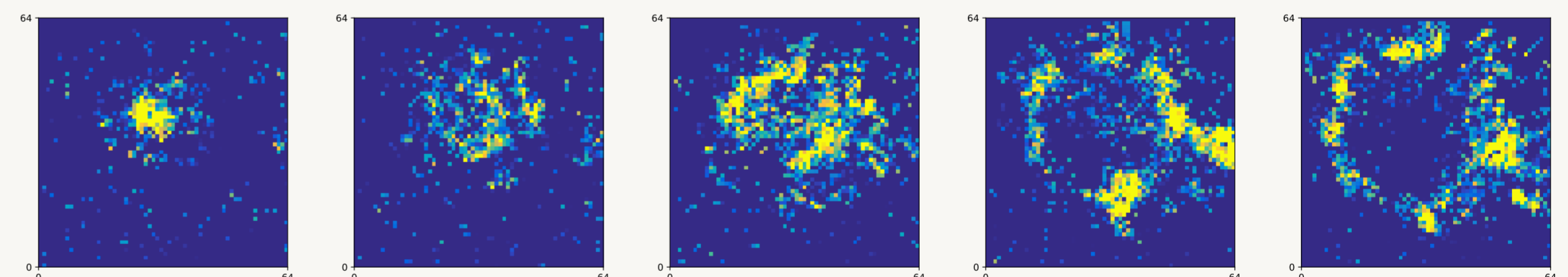
**Figure 4. Illustration of inner retina and recording setup.** Spontaneous retinal waves are generated in a layer of laterally interconnected amacrine cells. These waves activate Retinal Ganglion Cells (RGCs), the output cells of the retina. RGC electrical activity is recorded via a  $64 \times 64$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

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

- 4096-electrode arrays (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)

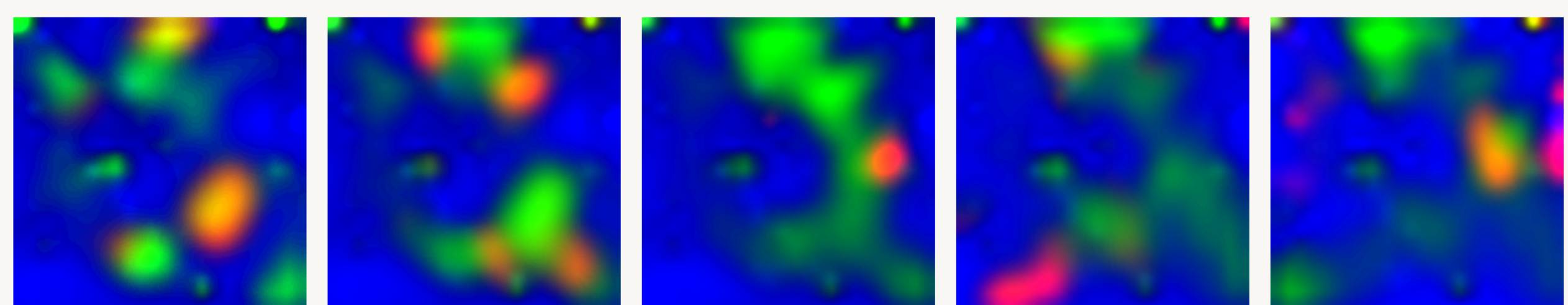


**Figure 5. 4096-electrode array.** Left: Array (3). Right: Spikes recorded in a single session.



**Figure 6. Example wave event, spike histograms in one-second intervals.** Mouse retina, postnatal day 11.

## 6. Bayesian filtering recovers latent states



**Figure 7. Filtering recovers retinal wave states.** Frames shown every 48 seconds; postnatal day 10.

### 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: optimize for fast parameter inference and apply to basic neuroscience