#### Reproducing kernel Hilbert Spaces for Spike Train Analysis

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## Outline



- Motivation and goals
- Generalized Cross-Correlation
- Instantaneous Cross-Correlation
- Results

#### Motivation



- Need a mathematical framework for the analysis of multiple spike trains.
- The fundamental concept for data analysis is the definition of an inner product.
- The mathematical structure needed for analysis is available in the form of Reproducing Kernel Hilbert Spaces (RKHS).

#### Motivation



- Cross-correlation is widely used for spike train analysis, but there are four main limitations:
  - 1. Traditionally, it is applied to binned spike trains
  - 2. It assumes stationarity (and ergodicity)
  - 3. To deal with non-stationarity the analysis is done in moving windows, with a tradeoff between temporal resolution and estimation accuracy
  - 4. Analysis only for pairs of spike trains.

#### Goals



- Define a Reproducing Kernel Hilbert Space (RKHS) based on ideas for cross-correlation, to remove or alleviate these limitations
  - Binning-free and data efficient estimation
  - High-temporal resolution exploring the spatial dimension of multi-electrode recordings and capable of cope with nonstationarity.

## What is Cross-Correlation?

Cross-correlation is an inner product

$$C_{AB}^{bin}[l] = \frac{1}{M} \sum_{n=1}^{M} N_A[n] N_B[n+l]$$

- The mapping into the RKHS is binning, which maps time to amplitude randomness, but loses information since time is quantized
- What is the role of binning?
- Can an inner product be defined without these limitations?

## **Generalized Cross-Correlation**

- Binning is an intensity estimator!
- Hence, using the intensity functions of the underlying point processes we can write a generalized cross-correlation (GCC),

$$C_{AB}(\theta) = E \left\{ \lambda_A(t) \lambda_B(t+\theta) \right\}$$
$$= \lim_{T \to \infty} \frac{1}{2T} \int_{-T}^T \lambda_A(t) \lambda_B(t+\theta) dt$$

- This is a functional inner product using the full resolution of the recordings
- But, it still assumes stationarity (and ergodicity)

#### Generalized Cross-Correlation Estimation from data

 To estimate this inner product, we start by estimating the intensity function using kernel smoothing

$$\hat{\lambda}_A(t) = \sum_{m=1}^{N_A} h(t - t_m^A)$$

• Substituting is yields the estimator

$$\hat{C}_{AB}(\theta) = \frac{1}{T} \sum_{m=1}^{N_A} \sum_{n=1}^{N_B} \kappa_\tau \left( t_m^A - t_n^B + \theta \right)$$

#### Generalized Cross-Correlation Advantages

- Cross-correlation is a special case of the GCC, in which spike times are quantized and a rectangular function is used for  $\kappa_{\tau}$
- Operates directly on the spike times, utilizing the full resolution of the recordings
- Takes advantage of the sparse nature of spike trains
- Smoothing by  $\kappa_{\tau}$  allows for multiscale analysis, between synchrony or firing rate

## **Instantaneous Cross-Correlation**



But what about high temporal resolution?
GCC still assumes (piecewise) stationary analysis
Solution: drop the expectation over time!

 Then, the instantaneous cross-correlation (ICC) is defined as

$$\tilde{c}_{AB}(t,\theta) = \hat{\lambda}_A(t)\hat{\lambda}_B(t+\theta).$$

## Instantaneous Cross-Correlation Online estimation

- Estimation is very simple, even online...
- For online estimation, consider

$$h(t) = (1/\tau) \exp[-t/\tau] u(t),$$

as the smoothing function.

• The estimated intensity function is

$$\hat{\lambda}_A(t) = \frac{1}{\tau} \sum_{\substack{t_m^A \le t}} \exp\left(-\frac{t - t_m^A}{\tau}\right) u(t - t_m^A).$$

# Instantaneous Cross-Correlation Approximating the GCC

•  $\tilde{c}_{AB}(t,\theta)$  is a stochastic approximation of the GCC,

$$\frac{1}{T} \int_0^\infty \tilde{c}_{AB}(t,\theta) dt = \frac{1}{T} \sum_{m=1}^{N_A} \sum_{n=1}^{N_B} \frac{1}{2\tau} \exp\left(-\frac{|t_m^A - t_n^B + \theta|}{\tau}\right)$$
$$= \frac{1}{T} \sum_{m=1}^{N_A} \sum_{n=1}^{N_B} \kappa_\tau \left(t_m^A - t_n^B + \theta\right)$$
$$= \hat{C}_{AB}(\theta),$$

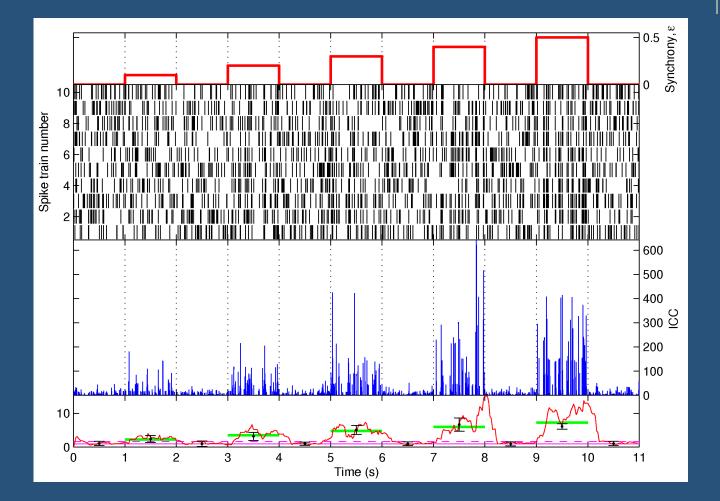
## Instantaneous Cross-Correlation As a neural ensemble measure

- To achieve high temporal resolution, the expectation over time was dropped, which the variance of the measurements...
- Solution: compute the expectation over the ensemble!

$$\bar{c}(t,\theta) = E\left\{\tilde{c}_{AB}(t,\theta)\right\}$$

 Giving rise to a spatio-temporal measure of correlations across the ensemble.

#### **Results** ICC as a synchronization measure



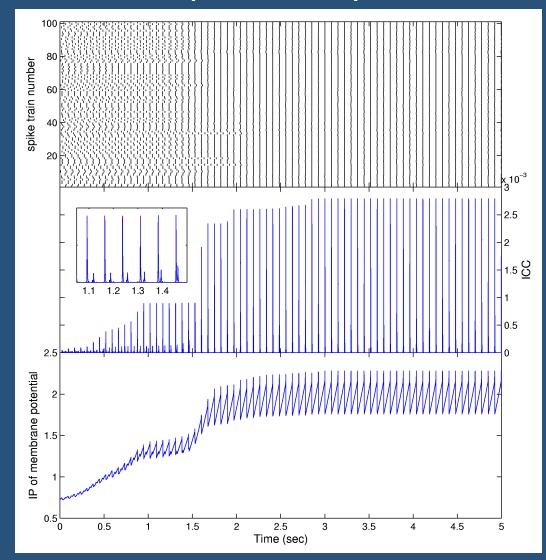


#### **Results** Synchronization of pulse-coupled oscillators



- Designed a spiking neural network of leaky integrate-and-fire neurons
- Network is known to synchronize over time (Mirollo & Strogatz, 1990)
- Utilized ICC to verify the evolution of synchrony
- Compared to an information-theoretic measure of internal coherence (membrane potentials)

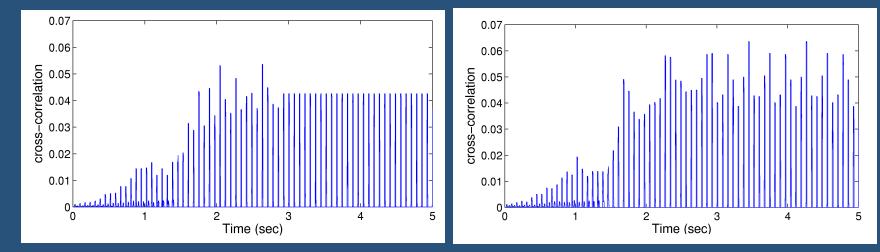
#### **Results** Synchronization of pulse-coupled oscillators



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#### Sensitivity of cross-correlation to bin size



#### Conclusion



- Utilized RKHS threory ideas to define GCC as an inner product of spike trains.
  - Computationally simple and accurate estimator
  - Natural extends towards multiscale analysis
- Suggested the ICC as a high temporal resolution measure of ensemble couplings.
  - Trades time averaging for "spatial" averaging
  - But, requires knowledge of groups of neurons over which to average

#### **Future work**



- Work on clustering and PCA of spike trains to find the ensembles of neurons over which to average the ICC.
- Most important of all, the RKHS allows for the development of more principled algorithms for computation with spike trains.