

# Spring project summary

Mark Blyth

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Discussion about single-cell and multi-cell approaches

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- ✦ Reuse Bath single-cell microfluidics device

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- ✂ Or could reuse Bath microfluidic device
  - ▶ Would require minor alterations to increase spatial resolution

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## Single- vs multi-cell

Deciding factors:

- ✖ No lab access for the foreseeable future
  - ▶ Work can be guided less by experiments
- ✖ Single-cell easier than multi-cell
  - ▶ I know enough about single-cell CBC to start working on it

Conclusion: work on single-cell case

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## Current goals

- ✦ Single-cell *in-silico* CBC
- ✦ Tutorial-review paper for numerical continuation



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## Challenges of *in-silico* CBC

Data aren't ideal to work with:

- ✶ Real signals are noise-corrupted
  - ▶ Difficult to filter off, since spikes contain lots of high-frequency components
  - ▶ Hard to run continuation on stochastic and noisy signals
- ✶ Neurons are fast-spiking
  - ▶ Fourier discretisation won't work
  - ▶ Discretisations need to be very high-dimensional, making Jacobian very slow to find

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    - ▶ Accurate derivatives, of arbitrary order
    - ▶ Allows for accurate delay embeddings, collocation discretisation – conventional continuation can be used on it
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 Truncated Fourier series

These require no preexisting knowledge [*loosely speaking*], and work well with sparse data

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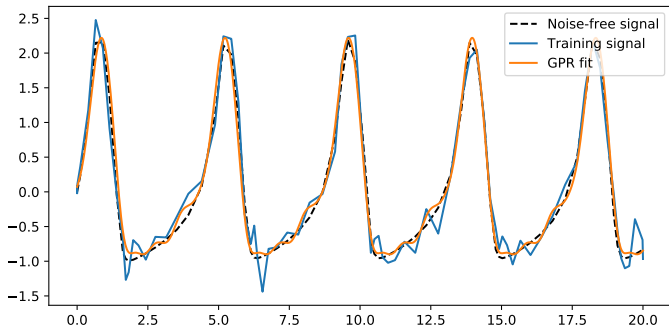
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  - ▶ Piecewise-polynomial model
- ✂ Gaussian process regression
  - ▶ Statistically optimal, when noise is Gaussian

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## Gaussian process regression

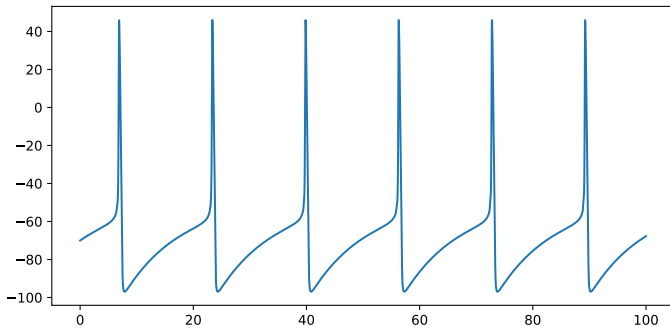


GPR can recover the underlying signal from noise-corrupted observations

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## Surrogate models

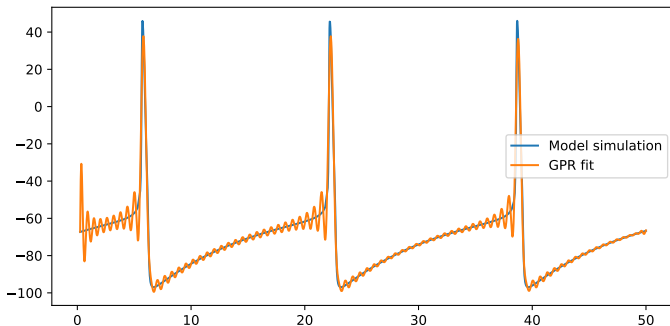


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- ✿ Most kernels are stationary
  - ▶ Assumes statistical properties are time-invariant *[they're not]*
  - ▶ Can't handle the spiking behaviours of neurons

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## Next questions

- ✦ Surrogate models on real data
- ✦ Predictor-corrector design
- ✦ Stochastic models

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## Continuation issues

- ✂ Discretisation is required to make predictor-corrector methods work
  - ▶ Can't run continuation on a function; must discretise it into a vector
- ✂ Discretisation has issues when used on fast-spiking data
  - ▶ Requires lots of datapoints
  - ▶ Slow to find a Jacobian for Newton-iterations
  - ▶ High noise-sensitivity
- ✂ Surrogate models and discretisation-free predictor-correctors might help overcome these

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## Alternative continuation approach

Predictor-corrector design:

- ✂ We could try discretisation-free predictor steps, using a surrogate model
  - ▶ Let  $f_i(t)$  be the surrogate model for system behaviours at parameter  $\lambda_i$
  - ▶ Given periodic orbits  $f_{i-1}$ ,  $f_i$ , predict  $f_{i+1} = f_i + h[f_i - f_{i-1}]$
- ✂ Corrector step would be harder



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## Stochastic models

Another challenge: real neurons are stochastic

- ✿ Stochasticity introduces new challenges
  - ▶ Coherence and stochastic resonance
  - ▶ Random attractors
  - ▶ Stochastic calculus
  - ▶ Not an area I know much about *[yet...]*
- ✿ Current work: CBC on noise-corrupted simulations
- ✿ Next work: CBC on truly stochastic models

Big question: how different would truly stochastic models be?

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## Goals

### Actions:

- ✂ Find a surrogate modelling method for neural data
- ✂ Attempt a discretisation-free corrector?
- ✂ Run CBC on deterministic models, then stochastic

### Results:

- ✂ Write up surrogate modelling into a conference abstract [July]
  - ▶ Maybe a conference paper [September]
- ✂ Use surrogate modelling for an *in-silico* CBC paper [next year?]

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 Functional kernel learning now works

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  - ✿ Support vector regression
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  - ▶ Gradient boosting: combine several weak learners to make a single strong learner

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