Inference for metapopulation dynamics

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Multiscale Microbial Communities
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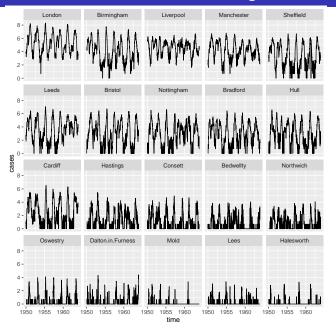
Slides are at https://ionides.github.io/talks/imsi22.pdf

Joint work with Kidus Asfaw, Ning Ning, Joonha Park and Aaron King

Inference challenges in population dynamics

- Combining measurement noise and process noise.
- Including covariates in mechanistically plausible ways.
- Continuous time models.
- Modeling and estimating interactions in coupled systems.
- Dealing with unobserved variables.
- Modeling spatial-temporal dynamics.
- Studying population dynamics via genetic sequence data.
- 1-6 are from Bjornstad & Grenfell (Science, 2001).
- 7 is from Grenfell et al (Science, 2004).
- 1–5 are largely solved, from a methodological perspective.

Example: Pre-vaccination measles in England & Wales



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Time series data, panel data & spatiotemporal data

- Looking at one unit (town) is time series analysis.
- Joint modeling of a few units (say, 2 or 3) is multivariate time series analysis.
- Analysis of many time series, without consideration of dynamic interactions, is panel data analysis.
- Allowing for coupling between units, we get **spatiotemporal analysis**, which in our context is **metapopulation analysis**.

Question: When should we avoid inference for spatiotemporal models? When do we need to consider coupling? How?

Desiderata

- We want to be able to fit arbitrary dynamic models. The limitations should be our scientific creativity and the information in the data.
- In practice, that means using plug-and-play methods which need a simulator from the model but not nice parametric equations.
- We want statistically efficient inference, to extract all the information in the data.
- In practice, that means using likelihood-based methods.
- In the time series case, iterated particle filtering (IF2) implemented in the R package pomp enables Masters-level statisticians to do this (https://ionides.github.io/531w22/). The science may be hard, but the statistics is becoming routine.

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Panel data

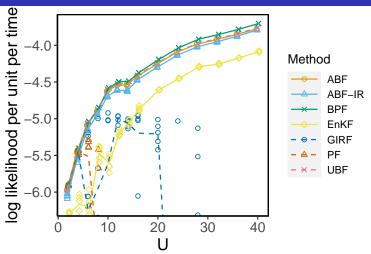
- To investigate epidemiological dynamics in multiple cities, one can consider each city independently, perhaps modeling a background immigration rate of infections for each city.
- **Decoupling** leads to panel data analysis, by assumption. Iterated filtering methods extend to panel data (Breto et al, *Journal of the American Statistical Association*, 2019).
- We must decide which parameters should be modeled as shared vs unit-specific.
- The consequences of decoupling are becoming easier to study with the development of statistical inference methods for coupled systems, i.e., metapopulation dynamics.

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The curse of dimensionality

- Particle filter (PF) methods are effective for inference on low-dimensional nonlinear partially observed stochastic dynamic systems. They scale exponentially badly.
- Extending the successes of particle filter methods from time series data to metapopulation data is becoming possible.
- Algorithms under consideration:
 Bagged filters (BF, IBF)
 Ensemble Kalman filter (EnKF, IEnKF)
 Guided intermiediate resampling filter (GIRF, IGIRF)
 Block particle filter (BPF, IBPF)
- Filters estimate latent states and evaluate the likelihood.
- Each filter has an iterated version which estimates parameters by repeated filtering using stochastic parameter perturbations.
- These algorithms are all implemented in an R package, spatPomp.

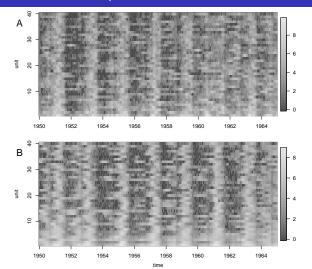
Filtering U units of a coupled measles SEIR model



Simulated data using a gravity model with geography, demography and transmission parameters corresponding to UK pre-vaccination measles (lonides et al, JASA, 2021).

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U=40 units for a coupled measles SEIR model



- **A**. Simulated Susceptible-Exposed-Infected-Recovered dynamics coupled with a gravity model (log of biweekly reported cases).
- B. Measles UK pre-vaccination case reports for the 40 largest cities.

More on the block particle filter

- BPF worked quickly, easily and reliably on our measles model filtering experiments.
- This motivated us to develop an IBPF for parameter estimation.
- BPF has theoretical support in some situations (Rebeschini & Van Handel, Annals of Applied Probability, 2015).
- BPF was independently proposed as the "factored particle filter" by Ng et al (2002, Proc. 18th Conference on Uncertainty and Artificial Intelligence) but not widely popularized.

Particle filter (PF)



Mutation

↓
Fitness

↓
tural solor

Natural selection

Particle filter algorithm

Predict: stochastic dynamics

Measurement: weight

Filter: resample

• PF is an evolutionary algorithm with good mathematical properties: an unbiased likelihood estimate and consistent latent state distribution.

Block particle filter (BPF)

Evolutionary analogy

Mutation ↓ Fitness for each chromosome ↓ Natural selection for each chromosome ↓ Recombine chromosomes

Block particle filter

Predict: stochastic dynamics

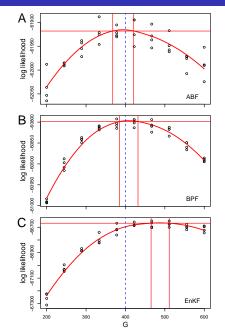
Measurement: weight for each block

Filter: resample for each block

Recombine blocks

- Blocks in BPF allow recombination (reassortment of chromosomes in sexual reproduction) in the evolutionary analogy.
- Blocks are a partition of the metapopulation units. Our experiments suggest treating each sub-population (i.e., city) as a block.

Measles likelihood slices for coupling parameter, G

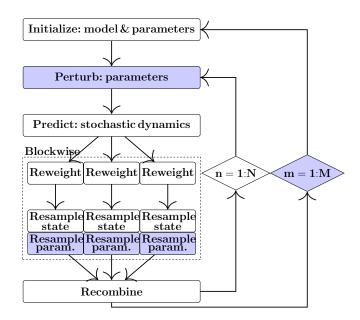


Simulating 15 year of data from U=40 cities for the measles model. Slice likelihood, varying G with other paramters fixed at the truth.

- **A**. Evaluation using adapted bagged filter (ABF).
- **B**. Evaluation using block particle filter (BPF).
- **C**. Evaluation using EnKF.

Results from lonides et al (2021, *JASA*). We computed a slice due to lack of good optimization algorithms to compute a profile.

An iterated block particle filter for parameter estimation



Scalability needed for practical inference

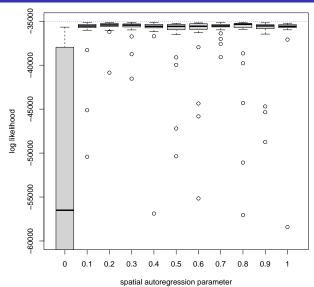
Large numbers of parameters

- Initial conditions will typically have to be estimated for each unit.
- Various dynamic parameters and measurement parameters (e.g., reporting rate) may also need to be unit-specific to obtain a statistical fit to the data.
- Working with hundreds of estimated parameters raises additional challenges on top of the high-dimensional coupled dynamics.

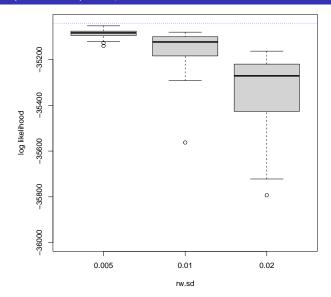
A moderate numbers of spatial units is enough to open new possibilities.

- As soon as dimension exceeds capabilities of a particle filter (say, U=5) we are in situations where likelihood-based inference for general models has been inaccessible.
- 10-100 coupled units is our target problem size.
- Larger problems will need numerical approximations (e.g., EnKF).
 Exact Monte Carlo methods help study the effect of these approximations.

Auto-regression of spatial perturbations for shared parameters



Random perturbations must be smaller to match larger number (20×13) of parameters



Likelihood on benchmark problems

	Simulated data	20 towns UK measles
Benchmark		
All parameters unit-specific		
4/13 parameters unit-specific		

- Simulated data: benchmark is likelihood at truth.
- Actual data: benchmark is likelihood from uncoupled model with all parameters unit-specific, and a parameter for immigration rate of new cases.

Future work

- We are getting close to the point where we can carry out likelihood-based inference for a flexible class of metapopulation models for measles. Flexibility supports generation and testing of scientific hypotheses.
- Measles was previously a motivating model system for POMP methods for single populations.
- Many systems in ecology, epidemiology and elsewhere could be studied in a SpatPOMP framework.

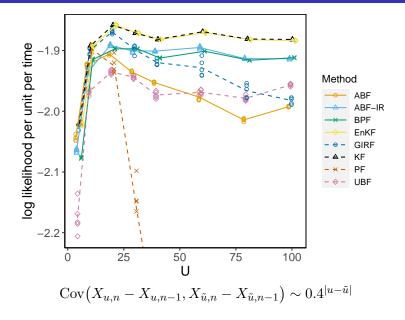
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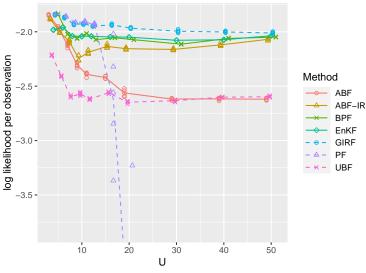
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Filtering *U*-dimensional correlated Brownian motion



Filtering U units of Lorenz 96 toy atmospheric model



$$dX_u(t) = \{X_{u-1}(t)(X_{u+1}(t) - X_{u-2}(t)) - X_u(t) + F\}dt + \sigma dB_u(t)$$