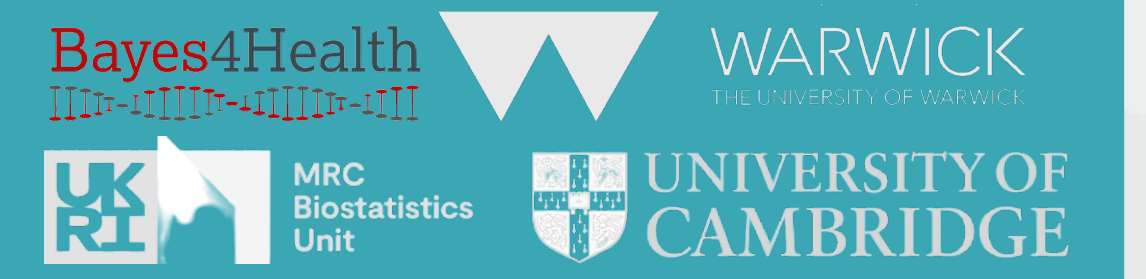


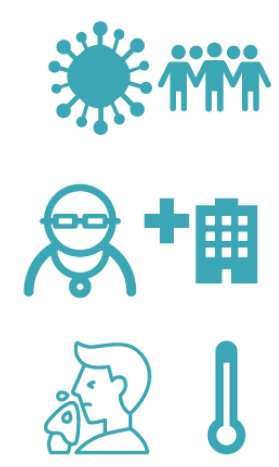
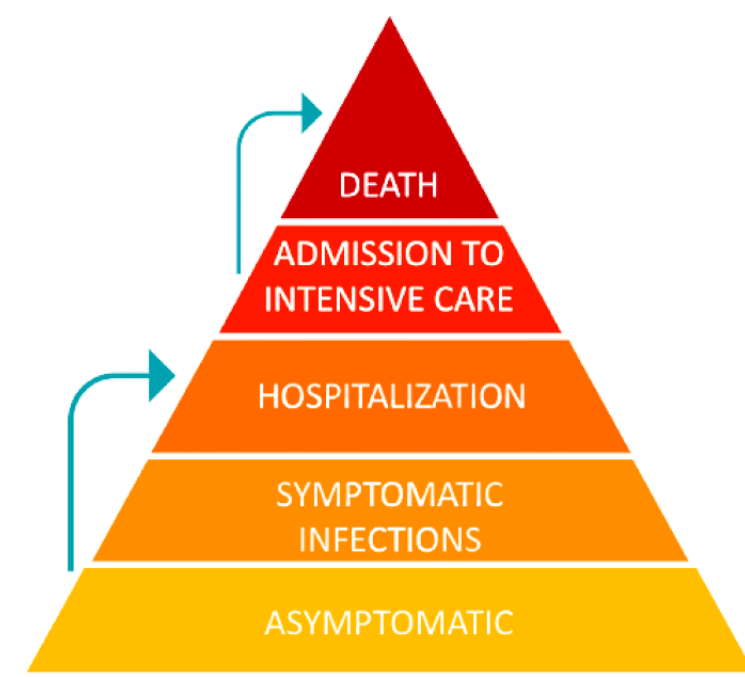
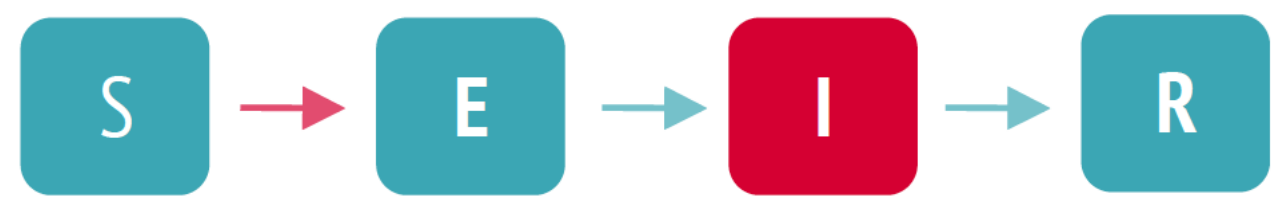
Inferring epidemics from multiple dependent data via pseudo-marginal methods

A Corbella, AM Presanis, PJ Birrell, and D De Angelis



Introduction: processes in epidemics

Multiple processes **jointly** contribute to generate epidemic data



Transmission dynamics in the population
Severity process for disease progression
Other sources of randomness

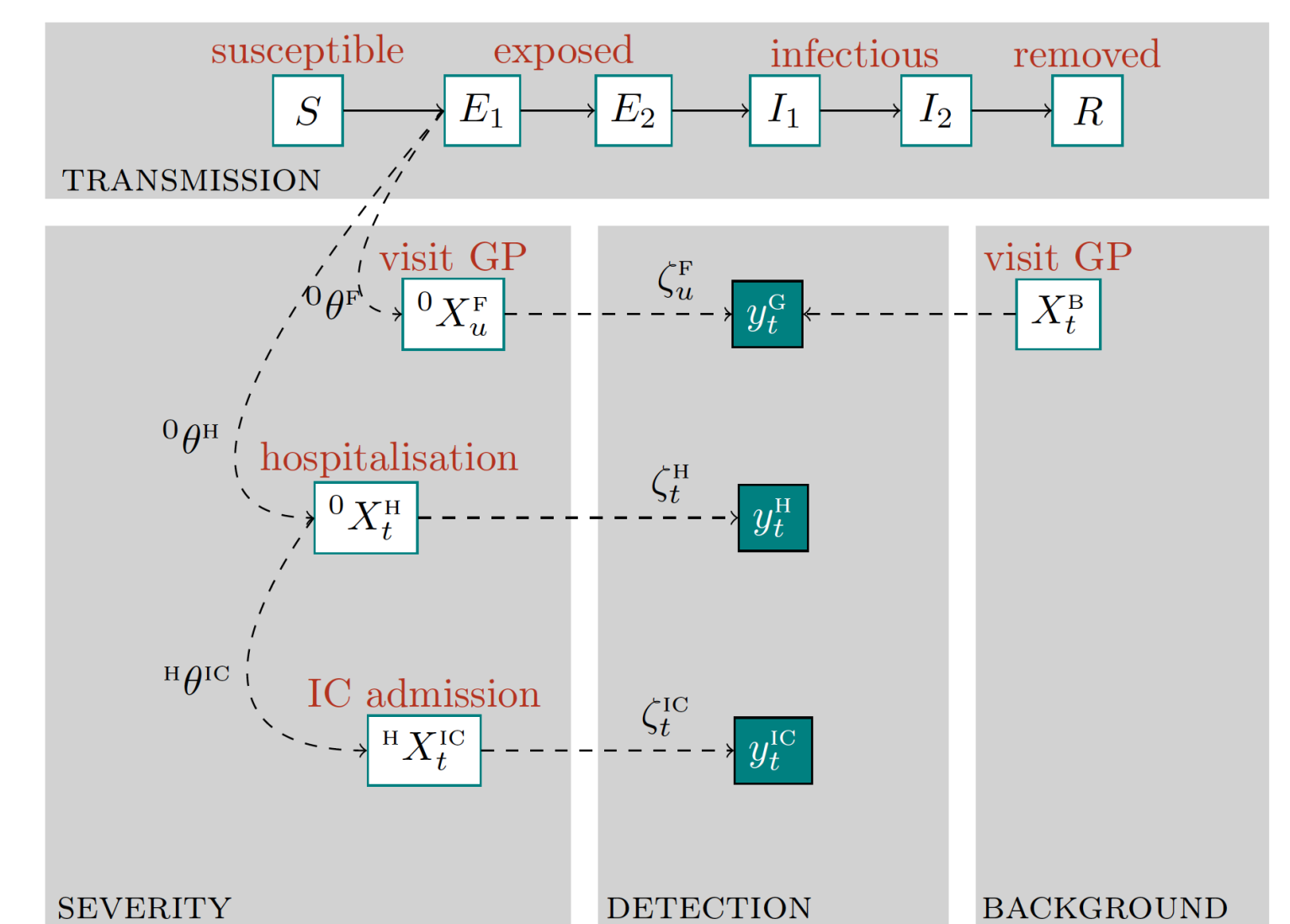
Which and how much **stochasticity** should we introduce?

A model for the 2017/18 Flu epidemic

Fuse data on

- weekly hospital and intensive care (IC) admissions
- daily GP consultations
- virology and serology

We assume a deterministic transmission model with **stochastic severity** as (1)



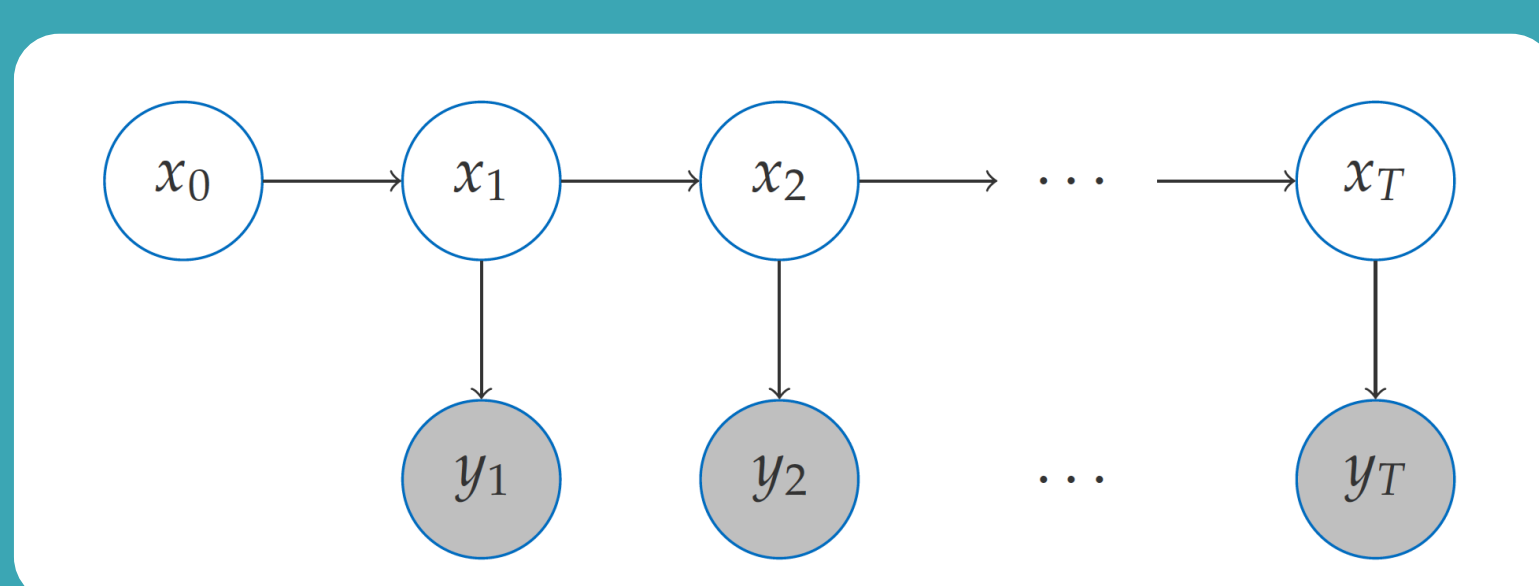
- background non flu ILLI cases:

$$\left(X_u^B \mid \mu_u^B\right) \sim \text{Pois}\left(\mu_u^B\right), \text{ with } \mu_u^B \text{ r.v. centred in } b_u = e^{\left\{\nu_1 + \nu_2 \cos\left(\frac{2\pi t u}{52}\right) + \nu_3 \sin\left(\frac{2\pi t u}{52}\right)\right\}}$$

- Accounts for discrete delay between events
- Day-of-the-week effect on the detection parameter $\omega_{[u]}$
- Virology data distribution

$$\left(Y_u^V \mid N_u^V, \mu_u^F, \mu_u^B\right) \sim \text{Binom}\left(N_u^V, \frac{\mu_u^F}{\mu_u^F + \mu_u^B}\right)$$

SMMs and Pseudo-Marginal methods



State Process
 $X_0 \mid \theta \sim p(x_0 \mid \theta)$
 $X_t \mid (X_{t-1}, \theta) \sim p(x_t \mid x_{t-1}, \theta)$
Observation process
 $Y_t \mid (X_t, \theta) \sim p(y_t \mid x_t, \theta)$

The likelihood can be computed **sequentially**

$$p(y_{1:T} \mid \theta) = \prod_{t=1}^T p(y_t \mid y_{1:t-1}, \theta) = \prod_{t=1}^T \int_{X_{0:t}} p(y_t, x_{0:t} \mid y_{1:t-1}, \theta) dx_{0:t}$$

Pseudo-Marginal methods consists of using a MC approximation of the likelihood $\hat{p}(y_{1:T} \mid \theta)$ within an MCMC routine.

Multiple data on epidemic

For large epidemics/pandemics a **semi-stochastic model** with deterministic transmission and stochastic severity is realistic

$$\left({}^0 X_t^H\right) \sim \text{Pois}\left({}^0 \xi_t \cdot {}^0 \theta^H\right)$$

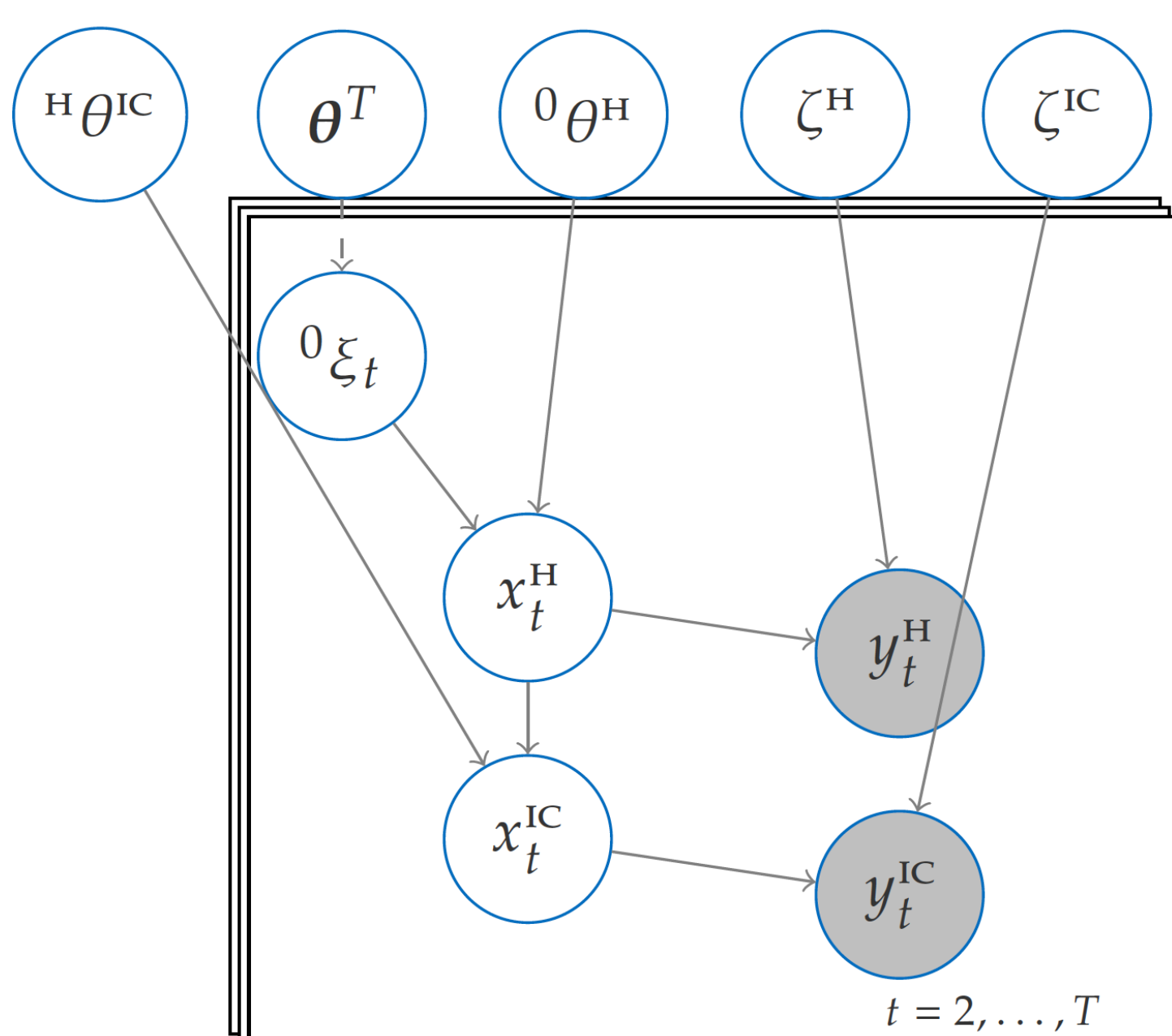
$$\left({}^0 X_{t+D}^H \mid {}^0 X_t^H = x_t^H\right) \sim \text{Multi}\left({}^0 x_t^H, {}^0 f_{0:D}^H\right), \quad X_t^H = \sum_{s=0}^S t-s {}^0 X_t^H$$

$$\left({}^H X_t^{IC} \mid X_t^H = x_t^H\right) \sim \text{Bin}\left(x_t^H, {}^H \theta^{IC}\right)$$

$$\left({}^H X_{t+D}^{IC} \mid {}^H X_t^{IC} = x_t^{IC}\right) \sim \text{Multi}\left({}^H x_t^{IC}, {}^H f_{0:D}^{IC}\right), \quad X_t^{IC} = \sum_{s=0}^S t-s {}^H X_t^{IC} \quad (1)$$

$$\left(Y_t^H \mid X_t^H = x_t^H\right) \sim \text{Bin}\left(x_t^H, \zeta_t^H\right)$$

$$\left(Y_t^{IC} \mid X_t^{IC} = x_t^{IC}\right) \sim \text{Bin}\left(x_t^{IC}, \zeta_t^{IC}\right)$$



Even if the temporal dependence is broken there is still **dependence in the severity states** so that:

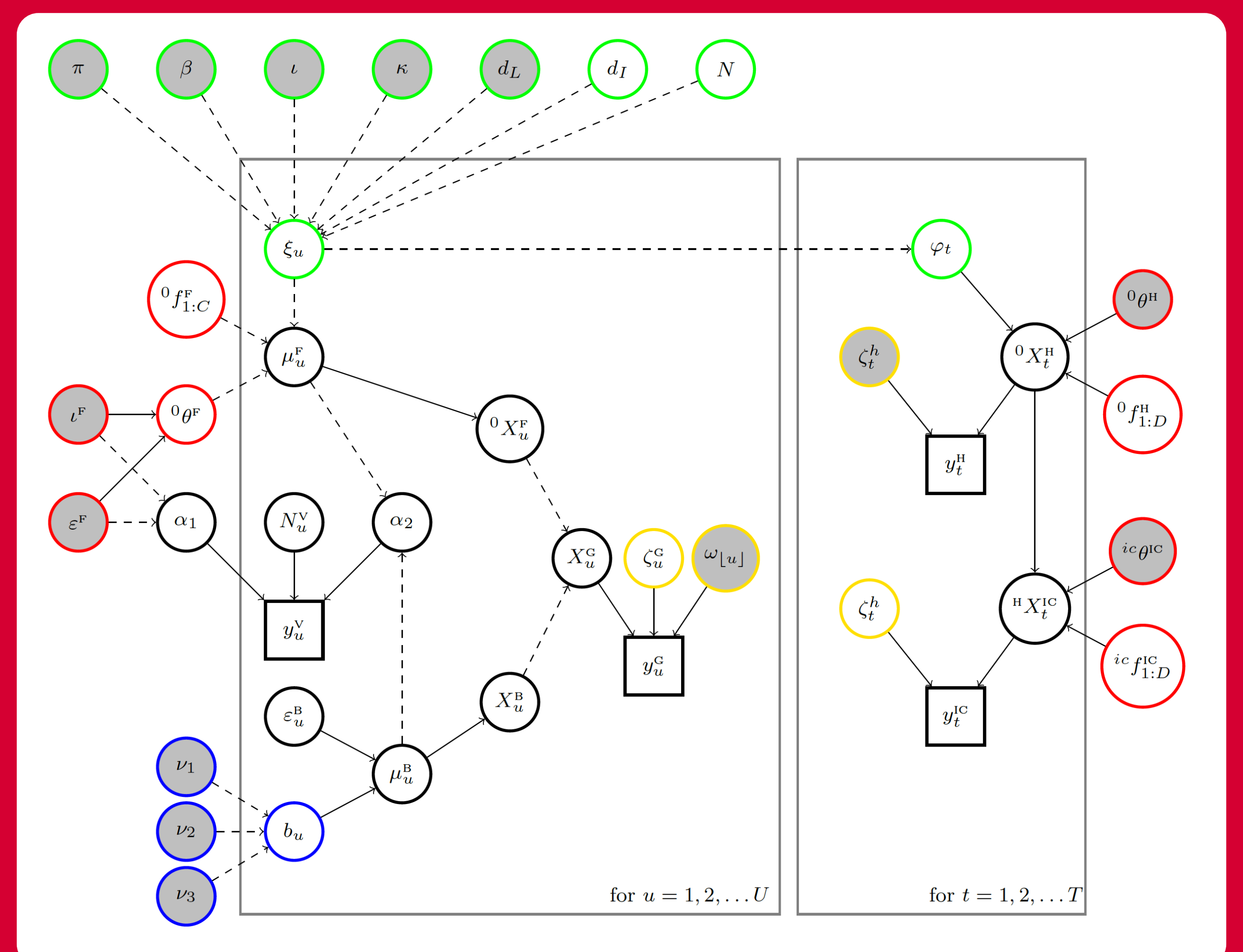
$$p(y_{1:T}, y_{1:T}^{IC} \mid \theta) \neq p(y_{1:T} \mid \theta) p(y_{1:T}^{IC} \mid \theta)$$

The joint likelihood factorises as:

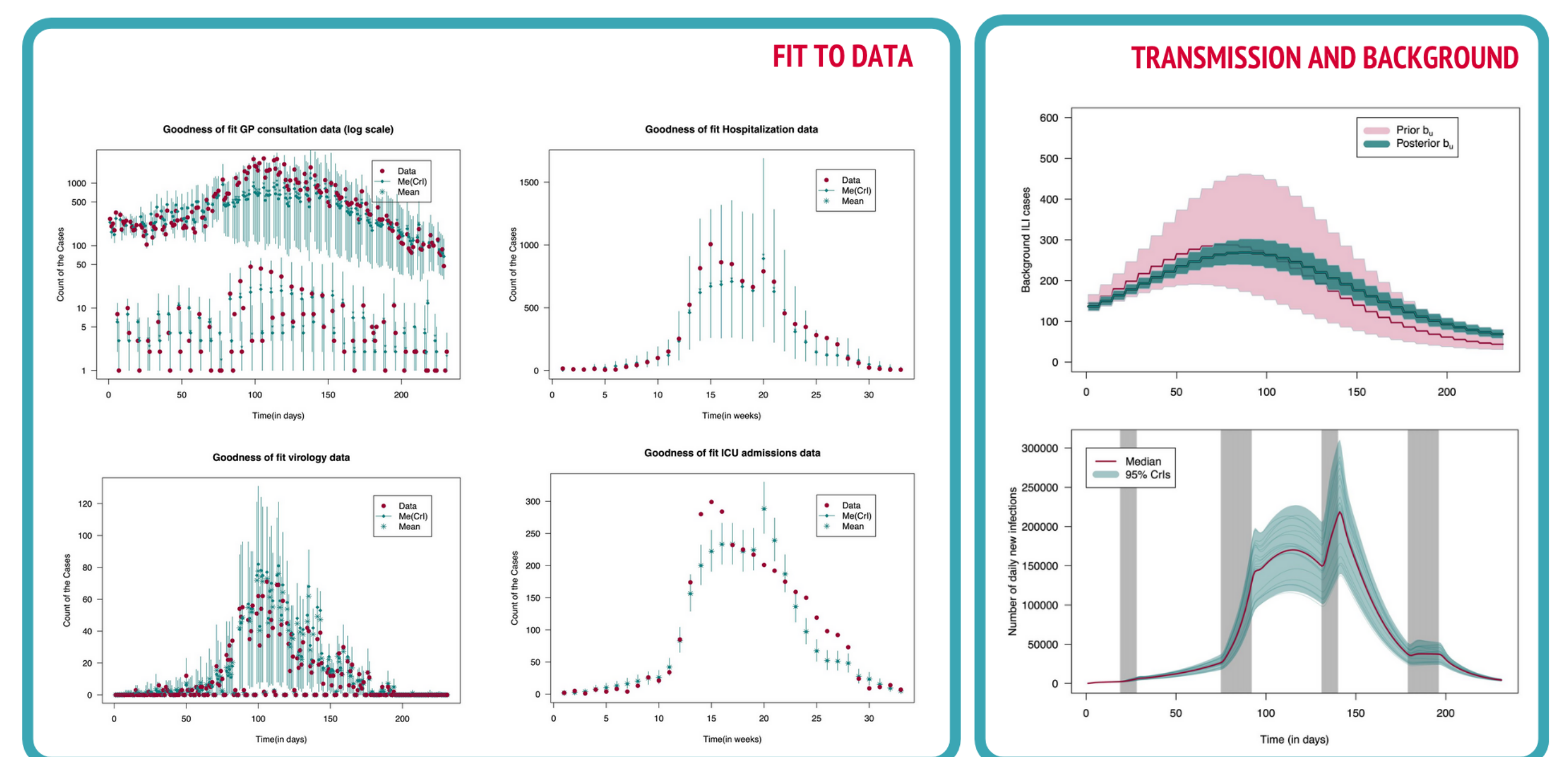
$$p(y_{1:T}, y_{1:T}^{IC} \mid \theta) = p(y_{1:T} \mid y_{1:T}^{IC}, \theta) p(y_{1:T}^{IC} \mid \theta) = p(y_{1:T} \mid y_{1:T}, \theta) p(y_{1:T} \mid \theta)$$

where one of the two components can be **approximated via MC integration** and embedded in a Pseudo-Marginal routine.

Model DAG



Results



Relevance of the dependence

Simulation study in a

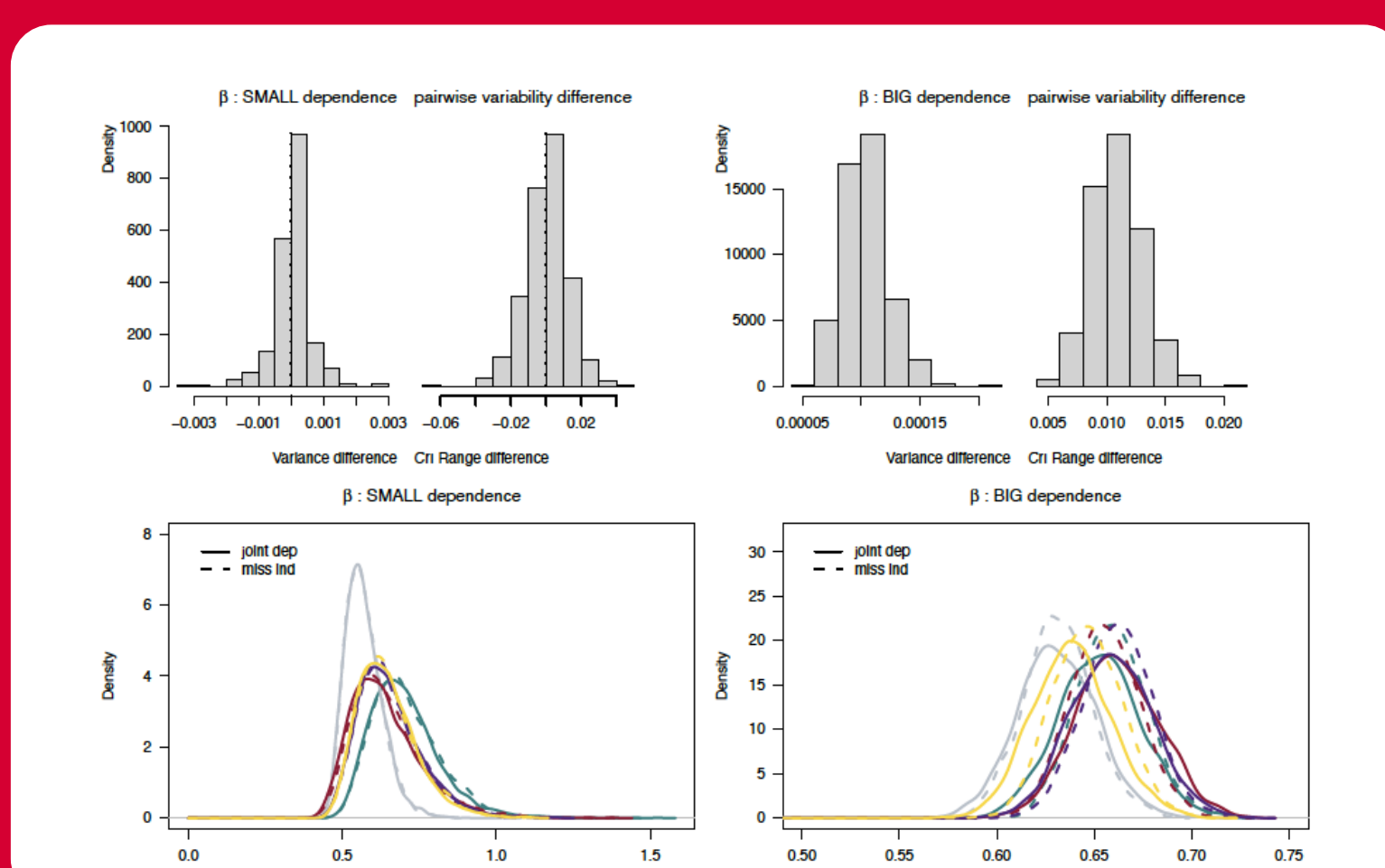
- small dependence scenario
- big dependence scenario

Compare the inference obtained with algorithms that assume

Misspecified independent

vs

Joint dependent data



Main Findings and Discussion

Findings

- general set up for epidemic inference as SMMs
- inferential tool for dependent data
- showed that it is key to account for dependence
- application to multiple dependent data on flu in 2017/18

What next?

- Improve the exploration of the parameter space with more efficient MCMCs
- Test the methods on models for prediction

Reference

Corbella A, Birrell PJ, Presanis AM and De Angelis D, **Inferring Epidemics from Multiple Dependent Data via Pseudo-Marginal Methods**. Submitted to *The Annals of Applied Statistics*.

