Introduction to ABC with an application to estimating transmission dynamics

Jukka Corander Department of Mathematics and statistics University of Helsinki, Finland





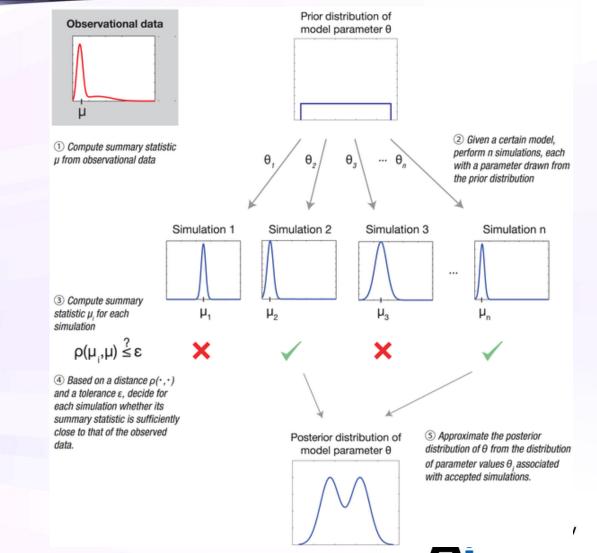
What ABC?

- Approximate Bayesian computation (ABC) is a method to do inference for intractable models
 Intractability means here that likelihood calculation is either too expensive or impossible
- Assumes we can still simulate data from our model
- •The core idea of ABC was introduced in a seminal paper by Tavaré et al. (1997) to do inference for a coalescence model





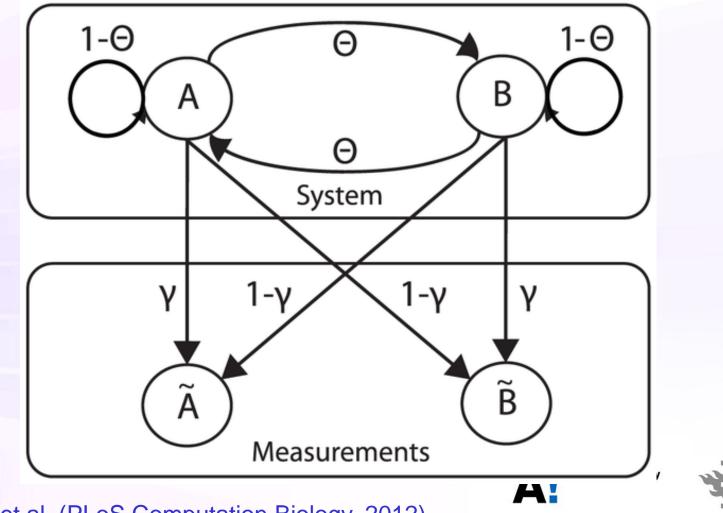
ABC - how does it work?



Sunnåker et al. (PLoS Computation Biology, 2012)

ABC – simple HMM example

Data: AAAABAABBAAAAAABAAAA, summary statistic #switches = 6



UNIVERSITY OF HELSINK

Sunnåker et al. (PLoS Computation Biology, 2012)

Assume prior $P(\theta) \sim U(0,1)$ and simulate data given random draws θ_i , *i*=1,...,*n*

| I | θ, | Simulated Datasets (Step 2) | Summary Statistic $\omega_{S^{\prime}}$ (Step 3) | Distance ρ (ω _{S,#} ω _d) (Step 4) | Outcome (Step 4) |
|---|------|-----------------------------|--|--|------------------|
| 1 | 0.08 | AABAAAABAABAAABAAAAA | 8 | 2 | accepted |
| 2 | 0.68 | AABBABABAAAABBABABBAB | 13 | 7 | rejected |
| 3 | 0.87 | BBBABBABBBBABABBBBBBA | 9 | 3 | rejected |
| 4 | 0.43 | AABAAAABBABBBBBBBBB | 6 | 0 | accepted |
| 5 | 0.53 | ABBBBBAABBABBABBABBBB | 9 | 3 | rejected |

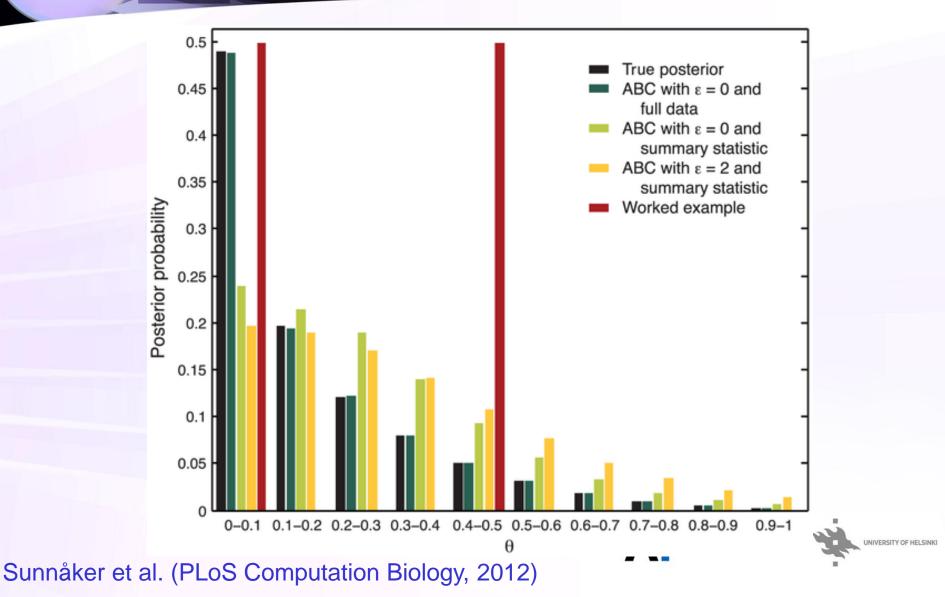
ABC – simple HMM example

doi:10.1371/journal.pcbi.1002803.t001

Sunnåker et al. (PLoS Computation Biology, 2012)



A look at inferences



ABC-MCMC

- 1. Sample candidate θ^* from proposal q(., θ) where θ is the current value of the parameter
- 2. Sample new data set using θ* and calculate new summary statistic S* (was S for θ)
- If ρ(S*,S)<ε, go to #4, else discard θ* and go to #1
- 4. Accept θ^* with probability $[\pi(\theta^*)/\pi(\theta)] \cdot [q(\theta,\theta^*)/q(\theta^*,\theta)]$
- 5. Return to #1

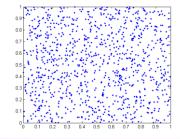




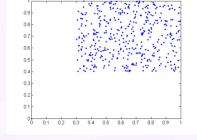




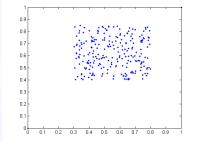


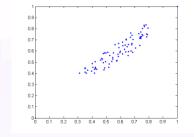


π(θ)



 $P(\theta|\rho(S(\theta),S) < \varepsilon_1)$





ABC-SMC

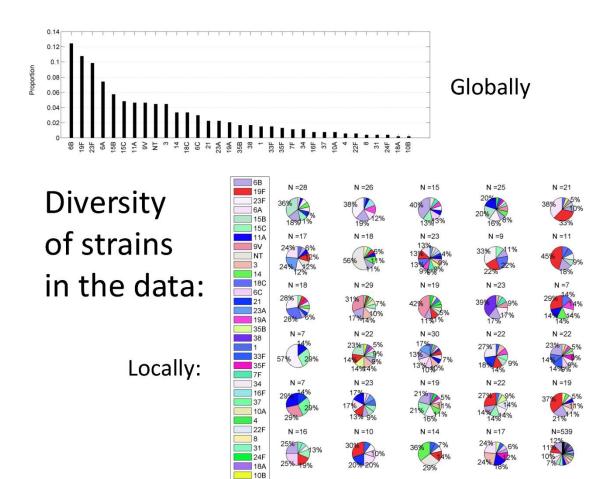
ABC recap

In reality more complex sampling algorithms: ABC-MCMC, particle filtering, etc
Necessitates quality controls, predictive checks,...

Formal ABC-based model comparison is an issue (Robert et al. PNAS, 2011), but latest results give more promising insight (Marin et al. JRSS B 2014, http://arxiv.org/abs/1110.4700)
Very intensive research area at the moment!



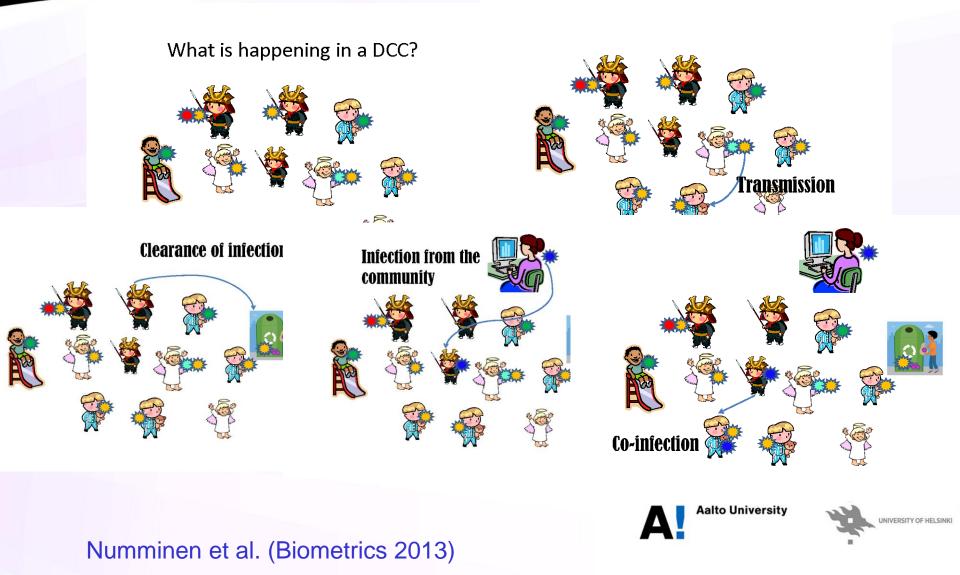
Pneumococcus strain incidences in Oslo DCCs (data sampled once in 2006)



NIVERSITY OF HELSINKI

Numminen et al. (Biometrics 2013)

Stochastic microepidemics in a metapopulation



Stochastic SIS-model for DCC transmissions

- Ingredients for a stochastic soup within a DCC:
- • $I_{ij}(t)$ indicator for kid *i* carrying strain *j* at time *t*
- β rate parameter for transmission from someone in DCC
- •Λ rate parameter for transmission from outside DCC
- θ competition parameter scaling the probability of coinfection
- γ clearance rate parameter, since we have data from single time point only, all other parameters are estimated relative to a fixed clearance rate



Stochastic SIS-model for DCC transmissions

Continuous-time Markov chain with transition probabilities:

$$Pr (I_{is}(t + \delta t) = 1 | I_{is}(t) = 0) = \beta E_s(I(t)) + \Lambda P^s + o(\delta t),$$

if $\sum_{j=1}^{N^s} I_{ij}(t) = 0$

$$Pr (I_{is}(t + \delta t) = 1 | I_{is}(t) = 0) = \theta (\beta E_s(I(t)) + \Lambda P^s) + o(\delta t)),$$

if $\sum_{j=1}^{N^s} I_{ij}(t) > 0$ and $I_{is} = 0.$

$$Pr (I_{is}(t + \delta t) = 0 | I_{is}(t) = 1) = \gamma + o(\delta t)$$
(2)

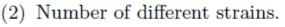


Aalto University

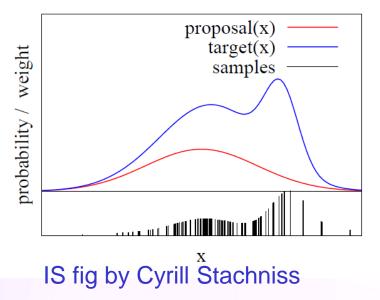
How to do the ABC inference?

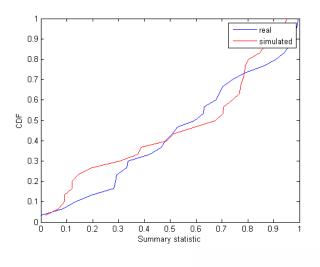
Summaries & discrepancies used in sequential importance sampling

- Shannon index of diversity (Peet, 1974) of the distribution of observed strains.
- $d_k = \int |F^k(x) \hat{F^k}(x)| dx.$



- (3) Prevalence of carriage among the observed individuals.
- (4) Prevalence of multiple infections among the observed individuals.







Some results

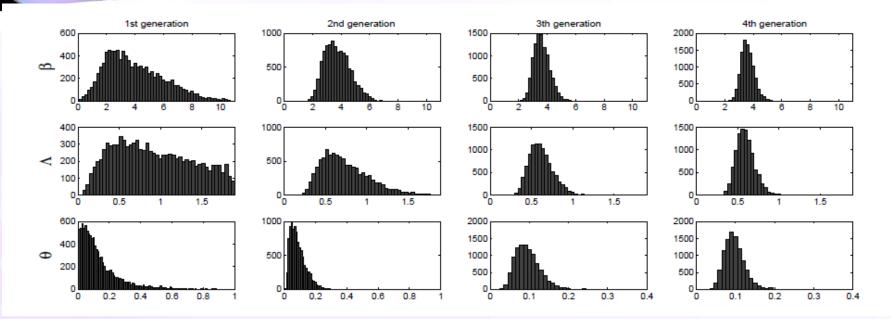


Table 1

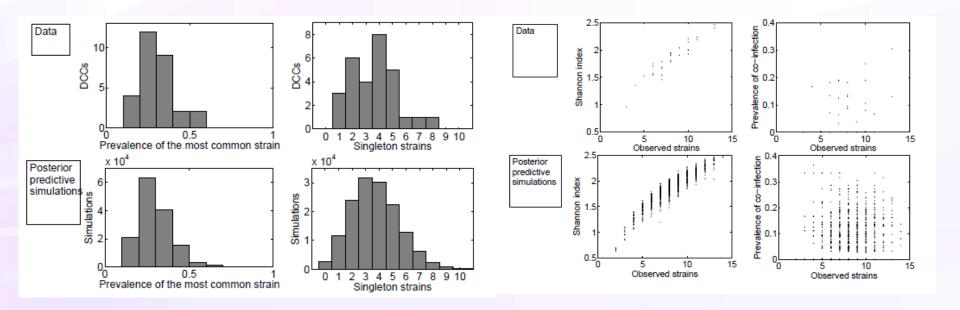
Summaries of the posterior distribution of the estimated parameters, with two different simulation times for the transmission model

| | Mean | Mean | 95% CI | 95% CI | | | |
|----------|--------|--------|-------------------|------------------|--|--|--|
| | T = 10 | T = 20 | T = 10 | T = 20 | | | |
| β | 3.589 | 3.594 | (2.8157, 4.5785) | (2.8113, 4.5621) | | | |
| Λ | 0.593 | 0.584 | (0.4017, 0.8359) | (0.3875, 0.8407) | | | |
| θ | 0.097 | 0.097 | (0.0605, 0.1422) | (0.0604, 0.1427) | | | |

Aalto University

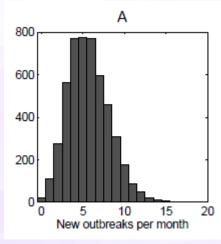


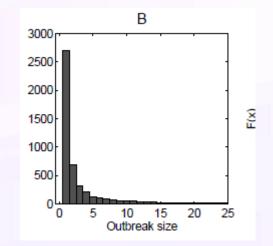
Model validation with 'unused' summaries

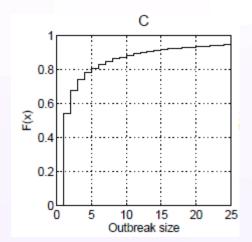


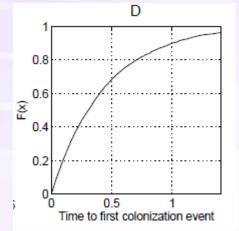


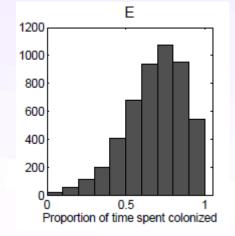
Posterior predictive simulations











Aalto University

What else with ABC?

- •ABC is particularly attractive for dynamic models with tricky/intractable/expensive likelihood functions
- ABC has grown particularly popular for complex spatio-temporal models in population genetics
 We are currently developing several generic machine learning inspired approaches to solve the key problems in ABC inference: choice of summary statistics, choice of metric to compare synthetic and real summaries, convergence to high likelihood/posterior regions





With great power comes great responsibility! -Uncle Ben

Hence the ABC sword should never be wielded casually



UNIVERSITY OF HELSIN