

# Bayesian analysis and antimicrobial resistance

Conor Rosato <sup>1</sup>

<sup>1</sup>University of Liverpool

December 14, 2023

# Overview

Background

Bayesian Analysis

# Background

- PhD titled “Disease Surveillance using Bayesian Methods”
- Six months into post-doc
- Working on antimicrobial resistance related problems

# Bayesian methods review

## Bayesian Calibration of Anti-microbial Resistance: A Review

CONOR ROSATO<sup>1</sup>, ALESSANDRO GERADA<sup>1</sup>, ALEX HOWARD<sup>1</sup>, JOHN HARRIS<sup>2</sup>, SIMON MASKELL<sup>3</sup>(Fellow, IEEE), and WILLIAM HOPE<sup>1</sup>,

<sup>1</sup>Department of Pharmacology and Therapeutics, University of Liverpool, 6 West Derby St, Liverpool, L7 7TX, United Kingdom

<sup>2</sup>Public Health England, UK

<sup>3</sup>Department of Electrical Engineering and Electronics, University of Liverpool, 9 Brownlow Hill, Liverpool, L69 3GB, United Kingdom

Corresponding author: Conor Rosato (e-mail: conor@liverpool.ac.uk)

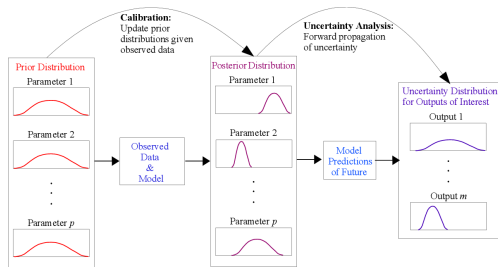
This research was funded in whole, or in part, by the Wellcome Trust (220691/Z/22/Z). For the purpose of Open Access, the author has applied a CC-BY public copyright license to any Author Accepted Manuscript version arising from this submission. CR, AH, JH and WH were funded by the Wellcome CAMO-Net UK grant: 225691/Z/22/Z and SM was funded by EPSRC through the Big Hypothesis under Grant EP/R01553/1.

**ABSTRACT** Antimicrobial resistance (AMR) emerges when disease-causing microorganisms develop the ability to withstand the effects of antimicrobial medicines. This phenomenon is often fuelled by the human-to-human transmission of pathogens and the overuse of antibiotics. Over the past 50 years, the increased computational power has facilitated the routine application of Bayesian inference algorithms, such as Markov Chain Monte Carlo (MCMC) and Sequential Monte Carlo (SMC) methods. These algorithms are instrumental in calibrating complex statistical models to vast amounts of data. In this comprehensive review, we delineate and exemplify the development and application of Bayesian methods within the context of AMR, offering practical examples. Additionally, we outline recent advancements in the methodology of these inference algorithms and provide insights into their prospective applicability for modelling AMR in the future.

- Population based AMR
  - Spread of MRSA in hospital setting
- En vitro studies
  - The Minimal Inhibitory Concentration (MIC) to establish the susceptibility or resistance of antibiotics when tested against antimicrobials
- Specialised areas of AMR
  - Molecular sequences associated with drug resistance using phylogenetic trees
  - Population pharmacokinetic models - depict relationships between patient characteristics and drug exposure

# Bayesian Calibration

- Finding a unique set of model parameters that provide a good description of the system behaviour
- Bayesian methods are naturally suited for calibration because they estimate the input parameter's posterior distributions



# Bayesian Calibration

- Advantages
  - Provides a framework for dealing with uncertainty
  - Include prior information when modelling
  - Distribution over any hypotheses rather than point estimates
- Disadvantages
  - Can be hard to select a correct prior
  - Computer time intensive

# Bayesian calibration

- Two sampling algorithms include: Markov Chain Monte Carlo and Sequential Monte Carlo methods

Sampling Algorithm	Proposal	Studies
MCMC		[9], [16], [17], [36], [41]
MCMC	MB	[10], [14], [23], [28], [29], [34], [39], [40], [48]–[50]
Adaptive MCMC	MB	[14], [35], [49]
MCMC	Gibbs	[22], [24], [32], [44], [46]–[48]
MCMC	Langevin	[32]
MCMC	NUTS	[27], [31], [39], [38], [45]
SMC Sampler	ABC proposal	[37]
Particle-MCMC	MB	[33], [51]
SMC <sup>2</sup>	MB	[30]

TABLE 2: Inference Algorithms

- PhD focused on being accurate and efficient
- Aiming to work on AMR related problems

## An $\mathcal{O}(\log_2 N)$ SMC<sup>2</sup> Algorithm on Distributed Memory with an Approx. Optimal L-Kernel

Conor Rosato<sup>a</sup>, Alessandro Varsi<sup>a</sup>, Joshua Murphy<sup>a</sup> and Simon Maskell<sup>b</sup>

<sup>a</sup> Department of Pharmacology and Therapeutics, University of Liverpool, United Kingdom  
<sup>b</sup> Department of Electrical Engineering and Electronics, University of Liverpool, United Kingdom  
 Email: {conor.ra, a.varsi, joshua.murphy, simonm}@liverpool.ac.uk

API 21 Nov 2023

**Abstract**—Calibrating statistical models using Bayesian inference often requires both accurate and timely estimates of parameters of interest. Particle Markov Chain Monte Carlo (pMCMC) and Sequential Monte Carlo Squared (SMC<sup>2</sup>) are two methods that use an unbiased estimate of the log-likelihood obtained from a particle filter (PF) to evaluate the target distribution. Full SMC<sup>2</sup> constructs a single Markov chain which is supported by a set of nodes to model distribution shifts. Distributed Memory (DM) architectures. This is in contrast to SMC<sup>2</sup> which involves processes, such as sequential sampling, that are described as end-to-endly parallel. However, difficulties arise when attempting to parallelise sampling. One of the main choices of backward kernel, recycling scheme and comput-

[1]. Theoretically, sampling with a single Markov chain using MCMC is accurate and ensures convergence to the correct posterior distribution. However, sampling with one MCMC chain cannot readily exploit parallel computing architectures such as Message Passing Interface (MPI). This is due to the fact MCMC samples being dependent on the current sample. Running multiple MCMC chains in parallel is a method for overcoming this issue [2]. The challenge of parallelising a single MCMC chain contrasts with SMC<sup>2</sup> samples which have inherent steps that can be described as end-to-endly parallel. Sampling  $N$  independent samples from a proposal

