## **Introduction to Statistical Modelling**

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#### Statistical inference



**Statistical inference** can be thought of the *inverse* of simulation.

That is, we observe some data and want to know:

What **parameter values** for a model produce the 'best fit' to the data?

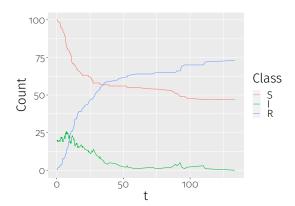
We can use this to provide insights into key epidemiological processes (e.g. e.g. estimating the transmission rate,  $R_0$  etc.). We can also use this to produce **predictions** and **forecasts**.

Key aspect is that we wish to quantify **uncertainty**.

### Example: SIR model



As an example, let's look at some data that we've simulated from a simple SIR model in a closed population of size N=120, with the introduction of 20 initial infectives at time t=0.





If we assume these data come from a **stochastic** SIR model of the form:

$$\begin{split} &P(S_{t+\delta t} \to S_t - 1 \text{ and } I_{t+\delta t} \to I_t + 1) \approx \beta S_t I_t, \\ &P(I_{t+\delta t} \to I_t - 1 \text{ and } R_{t+\delta t} \to R_t + 1) \approx \gamma I_t \end{split}$$

for small  $\delta t$ . We can then ask the question:

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This question can be tackled by appealing to the **likelihood function**.

The likelihood function,  $f(\mathbf{y} \mid \theta)$ , gives the **likelihood**<sup>†</sup> of observing the data  $(\mathbf{y})$  **given** a set of parameters  $(\theta)$ .

The exact form of the **likelihood** function depends on the **specific model** and **data**.

 $<sup>^{\</sup>dagger}$  if the data,  $\mathbf{y}$ , are **discrete**, then this is a **probability** 

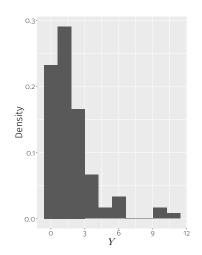


# The exact form of the **likelihood** function depends on the **specific model** and **data**.

For example, imagine we have n=100 independent samples from an **exponential** distribution:

$$Y_i \sim \text{Exp}(\lambda)$$

where  $\lambda$  is **unknown**.

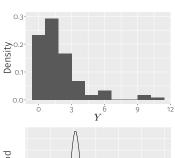


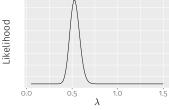


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If the data are **independent**, then

$$\begin{split} f(\mathbf{y} \mid \lambda) &= \prod_{i=1}^n f(y_i \mid \lambda) \\ &= \prod_{i=1}^n \lambda e^{-\lambda y_i} \end{split}$$



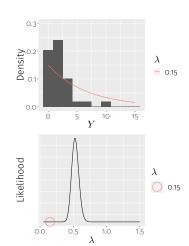




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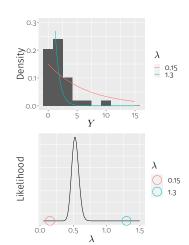




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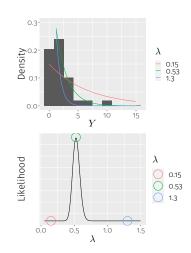




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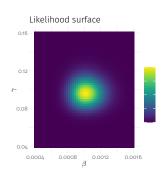




# The likelihood function can be thought of as a function of the unknown parameters $\theta$ .

In the case of our SIR model, we have  $\theta = (\beta, \gamma)$ .

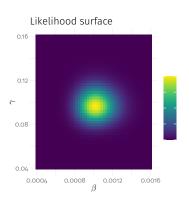
The **likelihood surface** (for different values of  $\beta$  and  $\gamma$ ) looks like the plot opposite.



Note that in general **likelihoods** for compartmental models like this are **intractable**<sup>†</sup>, but in this simulated setting we can write it down directly.

<sup>&</sup>lt;sup>†</sup>since **data** points are generally **not independent**, and typically the likelihood also depends on **unobserved variables**—we will return to this later





We can see that parameter values in the **yellow** region, produce **higher** likelihood values than parameter values in the **dark blue** regions.

This means that parameters in the **yellow** region would produce simulations that are **more consistent** with the observed data than parameters in the **dark blue** regions.

#### Maximum likelihood

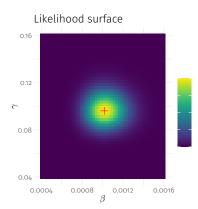


A natural way to estimate the parameters is to ask:

What parameter values **maximise** the likelihood function<sup>†</sup>?

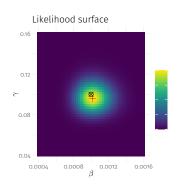
Here the **maximum likelihood** estimates are shown with a **red cross**, and are given by:

$$\hat{eta}=0.00102$$
 and  $\hat{\gamma}=0.0961,$  to 3 significant figures.



<sup>&</sup>lt;sup>†</sup>we will see an alternative approach—using the **Bayesian** framework—later





- The absolute value of the likelihood is rarely interpretable, only relative values.
- The likelihood is based on the data and the choice of model, and thus will change for different data sets and different models.
- ML estimates do not guarantee a good fit.
- Similar parameter values can give similar fits (uncertainty).

#### Confidence intervals



Uncertainties in the parameter estimates can be quantified using **confidence intervals**. **Wider** confidence intervals signal **larger** uncertainties.

Here 95% confidence intervals<sup>†</sup> are:

- $\beta$ : (0.000743, 0.00129)
- $\gamma$ : (0.074, 0.118)

**Note**: these do **not** correspond to a 95% probability that the true value is between the limits. Rather, it means that if the experiment were to be conducted an **infinite** number of times, 95% of the time the calculated CI would contain the true value<sup>‡</sup>.

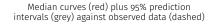
<sup>&</sup>lt;sup>†</sup>based on a **large sample** approximation

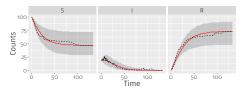
<sup>&</sup>lt;sup>†</sup>this is so-called **frequentist** inference, as opposed to **Bayesian** inference that we will cover shortly

### Model checking and prediction



We can check the model fit using the ML estimates to seed a large number of simulations from the model, and plot these against the observed data.





Here the model produces simulations that are consistent with the data<sup>†</sup>.

Note that the uncertainty bounds here **do not** account for the **parameter uncertainty**<sup>‡</sup>; to calculate a **true prediction interval** for these types of model is harder (see Gelman and Hill (2007) for *simulation-based* approaches).

<sup>†</sup>be careful, simulations from stochastic models can be tricky—see McKinley, Cook, and Deardon (2009)

<sup>\*</sup>the parameters are **fixed** at the MLEs

#### Practical



In the first practical we will explore fitting the **catalytic model** for endemic diseases to serology data for **rubella**.

To do this, we will need to write down a **likelihood function**, and then use one of R's in-build **optimisation** functions (optim()) to maximise with respect to the parameters to find the **maximum likelihood estimates**.

#### References i



Gelman, Andrew, and Jennifer Hill. 2007. Data Analysis Using Regression and Multilevel/Hierarchical Models. Cambridge University Press.

McKinley, Trevelyan J., Alex R. Cook, and Robert Deardon. 2009. "Inference in Epidemic Models Without Likelihoods." *The International Journal of Biostatistics* 5 (1). https://doi.org/10.2202/1557-4679.1171.