

## Lesson 2: Simulation of stochastic dynamic models

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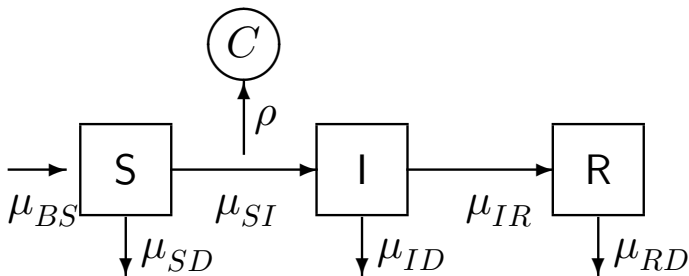
# Objectives

This tutorial develops some classes of dynamic models relevant to biological systems, especially for epidemiology.

1. Dynamic systems can often be represented in terms of *flows* between *compartments*.
2. We develop the concept of a *compartment model* for which we specify *rates* for the flows between compartments.
3. We show how deterministic and stochastic versions of a compartment model are derived and related.
4. We introduce Euler's method to simulate from dynamic models.
5. We specify deterministic and stochastic compartment models in pomp using Euler method simulation.

## A basic compartment model: The SIR model I

- ▶ We develop deterministic and stochastic representations of a susceptible-infected-recovered (SIR) system, a fundamental class of models for disease transmission dynamics.
- ▶ We set up notation applicable to general compartment models (Bretó et al. 2009).



## A basic compartment model: The SIR model II

S : susceptible

I : infected and infectious

R : recovered and/or removed

C : reported cases

- ▶ We suppose that each arrow has an associated rate, so here there is a rate  $\mu_{SI}(t)$  at which individuals in  $S$  transition to  $I$ , and  $\mu_{IR}$  at which individuals in  $I$  transition to  $R$ .
- ▶ To account for demography (births/deaths/migration) we allow the possibility of a source and sink compartment, which is not usually represented on the flow diagram. We write  $\mu_{BS}$  for a rate of births into  $S$ , and denote mortality rates by  $\mu_{SD}$ ,  $\mu_{ID}$ ,  $\mu_{RD}$ .
- ▶ The rates may be either constant or time-varying.
- ▶ For the simplest SIR model, ignoring demography, we set

$$\mu_{BS} = \mu_{SD} = \mu_{ID} = \mu_{RD} = 0.$$

## General notation for compartment models I

To develop a systematic notation, it turns out to be convenient to keep track of the flows between compartments as well as the number of individuals in each compartment:

- ▶  $N_{SI}(t)$ : the number of individuals who have transitioned from  $S$  to  $I$  **by** time  $t$ .  
We say that  $N_{SI}(t)$  is a *counting process*.
- ▶  $N_{IR}(t)$ : the number of individuals transitioning from  $I$  to  $R$  **by** time  $t$ .

To include demography, we could keep track of birth and death events by the counting processes:

- ▶  $N_{BS}(t)$ : the number of newborns into  $S$  **by** time  $t$ .
- ▶  $N_{SD}(t)$ ,  $N_{ID}(t)$ ,  $N_{RD}(t)$ : the number of deaths from  $S$ ,  $I$ , and  $R$  compartments **by** time  $t$ , respectively.

## General notation for compartment models II

- ▶ For discrete population compartment models, the flow counting processes are non-decreasing and integer valued.
- ▶ For continuous population compartment models, the flow counting processes are non-decreasing and real valued.

## Compartment model from counting processes

- ▶ The numbers of people in each compartment can be computed via these counting processes. Ignoring demography, we have:

$$S(t) = S(0) - N_{SI}(t)$$

$$I(t) = I(0) + N_{SI}(t) - N_{IR}(t)$$

$$R(t) = R(0) + N_{IR}(t)$$

- ▶ These equations represent *conservation of individuals* or *what goes in must come out*.

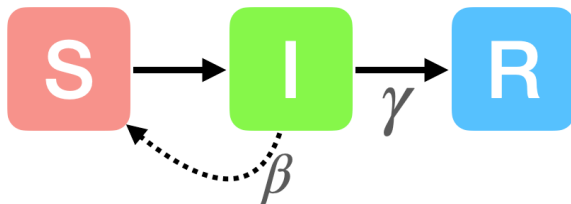
## Ordinary differential equation interpretation

Together with initial conditions specifying  $S(0)$ ,  $I(0)$  and  $R(0)$ , we just need to write down ordinary differential equations (ODEs) for the flow counting processes. These are:

$$\begin{aligned}\frac{dN_{SI}}{dt} &= \mu_{SI}(t) S(t) \\ \frac{dN_{IR}}{dt} &= \mu_{IR} I(t)\end{aligned}$$

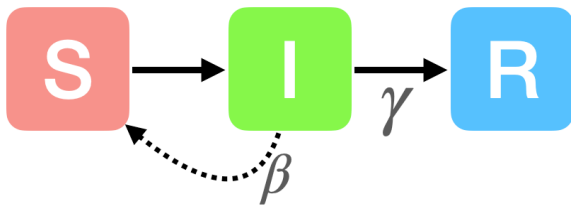


## Common notation for a deterministic SIR model



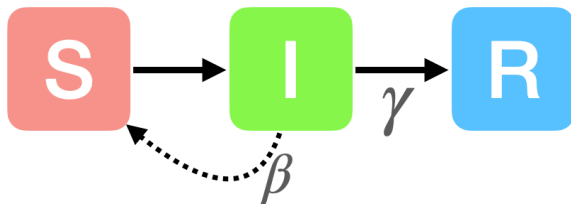
- ▶  $\beta$ : transmission rate, encompasses the frequency of contacts and transmission probability between individuals
- ▶  $\gamma$ : recovery rate, rate that infected individuals become “uninfectious”
  - ▶ Duration of infectiousness on average is  $\frac{1}{\gamma}$
- ▶  $S + I + R = N$

## Common notation for a deterministic SIR model - equations



$$\begin{aligned}\frac{dS}{dt} &= -\beta S \frac{I}{N} \\ \frac{dI}{dt} &= \beta S \frac{I}{N} - \gamma * I \\ \frac{dR}{dt} &= \gamma I\end{aligned}$$

## Common notation for a deterministic SIR model - Skeleton code



$$\begin{aligned}\frac{dS}{dt} &= -\beta S \frac{I}{N} \\ \frac{dI}{dt} &= \beta S \frac{I}{N} - \gamma * I \\ \frac{dR}{dt} &= \gamma I\end{aligned}$$

```
library(pomp)
Csnippet("
    DS = -Beta*S*I/N;
    DI = Beta*S*I/N - I*Gamma;
    DR = I*Gamma;
") -> sir_det_skel
```

# Stochastic Differential Equations (SDEs)

- ▶ By including randomness in the ODE system, we can have the stochastic differential equation (SDE) system.
- ▶ For example, for the ODE  $\frac{dx}{dt} = h(x)$ , a natural way to add stochastic variation is

$$\frac{dX}{dt} = h(X) + \sigma \frac{dB}{dt}$$

where  $\{B(t)\}$  is Brownian motion and so  $dB/dt$  is Brownian noise.

## The simple counting process and the reactions I

- ▶ A deterministic SIR model has a fixed trajectory, indicating that the number of each compartment at any time is fixed with given parameters and initial states; thus the transitions between compartments are fixed at any time.
- ▶ A stochastic SIR model, in the contrary, the trajectory and the transitions between compartments at any time are stochastic.
- ▶ Recall  $N_{SI}(t)$  and  $N_{IR}(t)$  are counting processes, indicating the number of total individuals transitioned from  $S$  to  $I$  and  $I$  to  $R$  by time  $t$ , respectively.
- ▶ A *simple counting process* is one which cannot count more than one event at a time.

## The simple counting process and the reactions II

- ▶ We then can relate the counting process to the common SIR reactions with the corresponding probabilities.
- ▶ Note that we are using *little o notation* and we write  $h(\delta) = o(\delta)$  to mean  $\lim_{\delta \rightarrow 0} \frac{h(\delta)}{\delta} = 0$ .

Table 1: Relationship between the counting processes, the reactions, and the probabilities.

Counting	Reaction	Probability
$N_{SI}(t + \delta) = N_{SI}(t) + 1$	$S \rightarrow S - 1$ $I \rightarrow I + 1$	$\beta S(t)I(t)\delta/N + o(\delta)$
$N_{SI}(t + \delta) = N_{SI}(t)$		$1 - \beta S(t)I(t)\delta/N + o(\delta)$
$N_{IR}(t + \delta) = N_{IR}(t) + 1$	$I \rightarrow I - 1$ $R \rightarrow R + 1$	$\gamma I(t)\delta + o(\delta)$
$N_{IR}(t + \delta) = N_{IR}(t)$		$1 - \gamma I(t)\delta + o(\delta)$

## The Euler's method I

- ▶ When referring the counting and its corresponding probability in Table 1, it is obvious that we can derive a continuous time Markov chain (CTMC) for the SIR model:

$$\begin{aligned}\mathbb{P}[N_{SI}(t + \delta) = N_{SI}(t) + 1] &= \beta S(t) I(t) / N \delta + o(\delta), \\ \mathbb{P}[N_{IR}(t + \delta) = N_{IR}(t) + 1] &= \gamma I(t) \delta + o(\delta).\end{aligned}$$

- ▶ For  $k = 1, 2, \dots$ , by discretizing this CTMC with small time step  $\delta$ , we can derive a numerical solution with the state variables  $\tilde{S}(k\delta)$ ,  $\tilde{I}(k\delta)$ ,  $\tilde{R}(k\delta)$ :

$$\begin{aligned}\tilde{S}(k\delta) &= S(0) - \tilde{N}_{SI}(k\delta) \\ \tilde{I}(k\delta) &= I(0) + \tilde{N}_{SI}(k\delta) - \tilde{N}_{IR}(k\delta) \\ \tilde{R}(k\delta) &= R(0) + \tilde{N}_{IR}(k\delta)\end{aligned}$$

- ▶  $\tilde{N}_{SI}(t)$  and  $\tilde{N}_{IR}(t)$ : the numerical solutions for  $N_{SI}(t)$  and  $N_{IR}(t)$

## The Euler's method II

- ▶ Let current  $t = k\delta$ , consider the small time interval  $t \leq \tau \leq t + \delta$ .
- ▶ Assume that the gradients  $\frac{dN_{SI}}{dt} = \mu_{SI}(t) S(t)$  and  $\frac{dN_{IR}}{dt} = \mu_{IR} I(t)$  are approximately constant
- ▶ We can have  $\tilde{N}_{SI}(t + \delta)$  and  $\tilde{N}_{IR}(t + \delta)$  as:

$$\begin{aligned}\tilde{N}_{SI}(t + \delta) &= \tilde{N}_{SI}(t) + \delta \beta S(t) I(t) / N \\ \tilde{N}_{IR}(t + \delta) &= \tilde{N}_{IR}(t) + \delta \gamma I(t)\end{aligned}$$



## The Euler's method III

Now we can include stochastic variation in the Euler's method.

- Recall the SDE:

$$\frac{dX}{dt} = h(X) + \sigma \frac{dB}{dt}$$

where  $\{B(t)\}$  is Brownian motion and so  $dB/dt$  is Brownian noise.

- An Euler approximation  $\tilde{X}(t)$  within the small time interval  $[t, t + \delta]$  and  $t = k\delta$  for  $k = 0, 1, 2, \dots$  is

$$\tilde{X}(t + \delta) = \tilde{X}(t) + \delta h(\tilde{X}(t)) + \sigma\sqrt{\delta} Z_k$$

where  $Z_1, Z_2, \dots$  are independent standard normal random variables, i.e.,  $Z_k \sim \text{Normal}(0, 1)$ .

- Although SDEs are often considered an advanced topic in probability, the Euler approximation doesn't demand much more than familiarity with the normal distribution.

## The Euler's method IV

Now we can consider applying the Euler's method for a stochastic SIR model:

- A binomial approximation with exponential transition probabilities.

$$\begin{aligned}\tilde{N}_{SI}(t + \delta) &= \tilde{N}_{SI}(t) + \text{Binomial}[\tilde{S}(t), 1 - \exp\{-\beta \tilde{I}(t)/N \delta\}], \\ \tilde{N}_{IR}(t + \delta) &= \tilde{N}_{IR}(t) + \text{Binomial}[\tilde{I}(t), 1 - \exp\{-\delta \gamma\}],\end{aligned}$$

where  $\text{Binomial}(n, p)$  is a binomial random variable with mean  $np$  and variance  $np(1 - p)$ . Here,  $p = 1 - \exp\{-\beta \tilde{I}(t)/N \delta\}$  and  $p = 1 - \exp\{-\delta \gamma\}$ , respectively.

## The Euler's method V

The following are two other ways for a stochastic SIR model with the Euler's approximation, what they are not as good as the previous one?

1. A Poisson approximation.

$$\tilde{N}_{SI}(t + \delta) = \tilde{N}_{SI}(t) + \text{Poisson}[\beta \tilde{S}(t) \tilde{I}(t)/N \delta],$$

where  $\text{Poisson}(\mu)$  is a Poisson random variable with mean  $\mu = \beta \tilde{S}(t) \tilde{I}(t)/N \delta$ .

2. A binomial approximation,

$$\tilde{N}_{SI}(t + \delta) = \tilde{N}_{SI}(t) + \text{Binomial}[\tilde{S}(t), \beta \tilde{I}(t)/N \delta].$$

## The Gillespie method I

- ▶ Numerical methods, such as the Euler's method, are approximations to the process by discretizing time using small time step  $\delta$
- ▶ However, the Gillespie method is the exact **Stochastic Simulation Method**, which leverages the Markov Property as well.
- ▶ In Table 1, by consider the reactions and the probabilities, we can derive the Gillespie algorithm for the stochastic SIR model.

## The Gillespie method II

With initialization,  $S(0)$ ,  $I(0)$ , and  $R(0)$ , at current time  $t$ :

1. Compute the total event rates:  $\lambda_1 = \beta S(t) I(t)/N$ ,  $\lambda_2 = \gamma I(t)$ ,  $\lambda = \lambda_1 + \lambda_2$
2. Compute the waiting time  $\Delta t \sim \text{Exponential}(\lambda)$
3. Select the reactions by sampling from probabilities  $(\frac{\lambda_1}{\lambda}, \frac{\lambda_2}{\lambda})$
4. Update the states from the selected reaction and update the time  $t \rightarrow t + \Delta t$
5. Repeat 1-4 till the end of the simulation time

Even though the Gillespie is an exact stochastic simulation method, it has limitations such as:

- ▶ Computational Intensity: For complex systems with many reactions, the Gillespie method can become computationally expensive.
- ▶ Rare Events: For systems where some reactions are very rare, a large number of simulation steps may be needed to capture these events, making the method slow.

## Euler vs. Gillespie

- ▶ Why and When would you prefer an implementation of Gillespie's algorithm to an Euler solution?

### Worked solution to the Exercise

- ▶ Numerically, Gillespie's algorithm is often approximated using so-called tau-leaping methods. These are closely related to Euler's approach. In this context, the Euler method has sometimes been called tau-leaping.

## Compartment models in pomp: The Consett Measles outbreaks I

Let's look at outbreak of measles in the town of Consett in England in 1948:

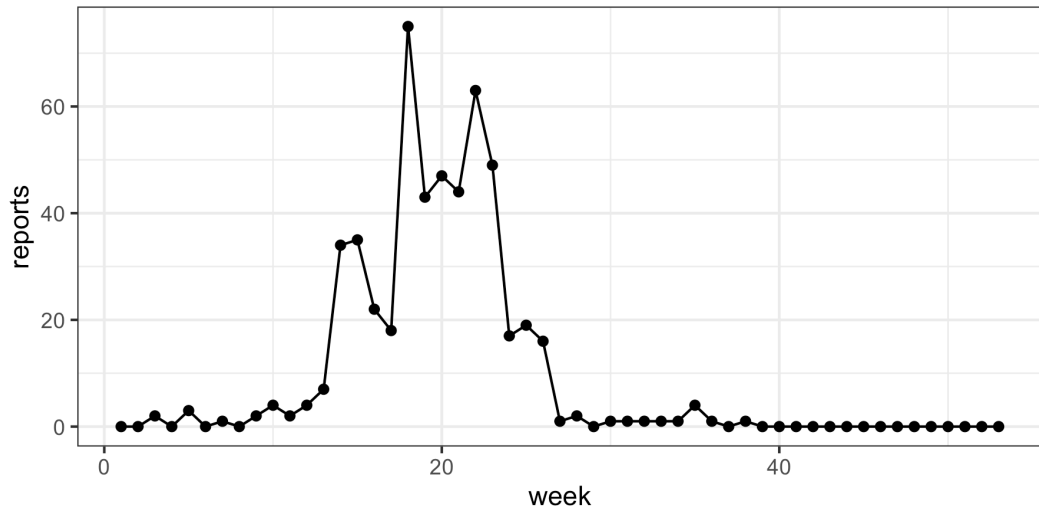
- ▶ the town had population of 38820,
- ▶ with 737 births over the course of the year.

```
library(tidyverse)
read_csv(paste0("https://kingaa.github.io/sbied/stochsim/",
  "Measles_Consett_1948.csv")) |>
  select(week, reports=cases) -> meas
meas |> as.data.frame() |> head(n=3)
```

week	reports
1	0
2	0
3	2

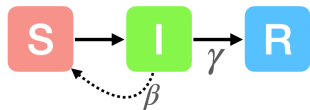
- ▶ week: time, indicates that the data are counted weekly
- ▶ reports variable: incidence, counts the number of reports of new measles cases each week

## Compartment models in pomp: The Consett Measles outbreaks II





## The SIR as a POMP model for measles I



- ▶ The **unobserved states**:  $S(t)$ ,  $I(t)$ ,  $R(t)$ , the numbers of individuals in the S, I, and R compartments, respectively.
- ▶ The constant population size:  $N = S(t) + I(t) + R(t)$ , as fixed at the known population size of 38,000.
- ▶ **Flows** move from one compartment to another over any particular time interval are modeled as *stochastic processes*.
- ▶ **Demographic stochasticity**: each individual in a compartment at any given time faces the same risk of exiting the compartment; the unavoidable randomness that arises from chance events occurring in a discrete and finite population.

## The SIR as a POMP model for measles II

Recall the application of the Euler's method to a stochastic SIR model.

- $\Delta N_{SI}$  and  $\Delta N_{IR}$ : the flows from S to I and from I to R over interval  $\Delta t$ , respectively:

$$\Delta N_{SI} \sim \text{Binomial} \left( S, 1 - e^{-\beta \frac{I}{N} \Delta t} \right),$$

$$\Delta N_{IR} \sim \text{Binomial} \left( I, 1 - e^{-\gamma \Delta t} \right).$$

## The SIR as a POMP model for measles III

- Implement the dynamics in pomp as an R function:

```
sir_stoch <- function (S, I, R, N, Beta, Gamma, delta.t, ...) {  
  dN_SI <- rbinom(n=1,size=S,prob=1-exp(-Beta*I/N*delta.t))  
  dN_IR <- rbinom(n=1,size=I,prob=1-exp(-Gamma*delta.t))  
  S <- S - dN_SI  
  I <- I + dN_SI - dN_IR  
  R <- R + dN_IR  
  c(S = S, I = I, R = R)  
}
```

- Note that, for a deterministic SIR model:

```
dN_SI <- Beta*S*I/N*delta.t  
dN_IR <- Gamma*I*delta.t
```

## The SIR as a POMP model for measles IV

We can implement the initialization function with the following assumptions:

- ▶ Assume the dynamics starts at week 0,  $t_0 = 0$ .
- ▶ At  $t_0$ , assume the initial number of infection is 1, that is  $I = 1$ .
- ▶ The initial number of susceptible is unknown, so we'll treat this fraction,  $\eta$ , as a parameter to be estimated.

```
sir_rinit <- function (N, Eta, ...) {  
  c(S = round(N*Eta), I = 1, R = round(N*(1-Eta)))  
}
```

## The SIR as a POMP model for measles V

With the initialization function `sir_rinit` and the process function `sir_stoch`, we can build a pomp object with these two components and the data:

```
library(pomp)
meas |>
  pomp(
    times="week", t0=0,
    rprocess=euler(sir_stoch, delta.t=1/7),
    rinit=sir_rinit
  ) -> measSIR
```

► Question: what do `times="week"` and `delta.t=1/7` indicate?

## The SIR as a POMP model for measles VI

- ▶ Assume the **observations**, the reports, result from a process by which new infections are diagnosed in a hospital and reported with probability  $\rho$ .
- ▶ The diagnosed infections are immediately hospitalized, therefore, they have, presumably, a much lower transmission rate; let's assume each *week's* reports as being related to the number of individuals who have moved from I to R over the course of that week.
- ▶ We then define a new variable,  $H$ , that tracks these daily counts.

## The SIR as a POMP model for measles VII

We now can modify the R functions to incorporate the new variable  $H$ :

```
sir_stoch <- function (S, I, R, N, Beta, Gamma, delta.t, H, ...) {  
  dN_SI <- rbinom(n=1,size=S,prob=1-exp(-Beta*I/N*delta.t))  
  dN_IR <- rbinom(n=1,size=I,prob=1-exp(-Gamma*delta.t))  
  S <- S - dN_SI  
  I <- I + dN_SI - dN_IR  
  R <- R + dN_IR  
  H <- H + dN_IR  
  c(S = S, I = I, R = R, H = H)  
}  
  
sir_rinit <- function (N, Eta, ...) {  
  c(S = round(N*Eta), I = 1, R = round(N*(1-Eta)), H = 0)  
}
```

## The SIR as a POMP model for measles VIII

Note that, we are so far accounting for the *flows* between compartments by days, while the reports are weekly cases. Since we want  $H$  to tally only the incidence over the week, we'll need to reset it to zero at the beginning of each week. Thus, in pomp terminology,  $H$  is an **accumulator variable**. We accomplish this using the `accumvars` argument to `pomp` when build the object:

```
measSIR |>
  pomp(
    rprocess=euler(sir_stoch,delta.t=1/7),
    rinit=sir_rinit,
    accumvars="H"
  ) -> measSIR
```

- ▶ Question: what does that mean by running a `pomp` function with the `pomp` object `measSIR`?



## The SIR as a POMP model for measles IX

Last but not least, we need to define a **measurement model** to relate the **observations**, reports, to the **unobserved** accumulative state,  $H$ .

- We will model the data by a negative binomial variable,

$$\text{reports}_t \sim \text{NegBin}(\rho H(t), k).$$

with mean  $\rho H(t)$  and variance  $\rho H(t) + (\rho H(t))^2/k$ . The binomial distribution does not have a separate variance parameter.

## The SIR as a POMP model for measles X

- To include the observations in the model, we must write either a `dmeasure` or an `rmeasure` component, or both:

```
sir_dmeas <- function (reports, H, Rho, k, log, ...) {  
  dnbinom(x=reports, size=k, mu=Rho*H, log=log)  
}  
  
sir_rmeas <- function (H, Rho, k, ...) {  
  c(reports=rnbinom(n=1, size=k, mu=Rho*H))  
}
```

## The SIR as a POMP model for measles XI

Eventually, we can add these two components to the previous measSIR object to update the dmeasure and rmeasure arguments:

```
measSIR |>
  pomp(
    rmeasure=sir_rmeas,
    dmeasure=sir_dmeas
  ) -> measSIR
```

## Specifying model components using C snippets I

- ▶ Although we can always specify basic model components using R functions, as above, we'll typically want the computational speed-up that we can obtain only by using compiled native code.
- ▶ pomp provides a facility for doing so with ease, using *C snippets*.
- ▶ C snippets are small pieces of C code used to specify basic model components.

## Specifying model components using C snippets II

- For example, a C snippet encoding the rprocess for an sir model is as follows.

```
sir_stoch <- Csnippet("  
  double dN_SI = rbinom(S,1-exp(-Beta*I/N*dt));  
  double dN_IR = rbinom(I,1-exp(-Gamma*dt));  
  S -= dN_SI;  
  I += dN_SI - dN_IR;  
  R += dN_IR;  
  H += dN_IR;  
")
```

## Specifying model components using C snippets III

### Note:

- ▶ It is necessary to define the data type for the real values `dN_SI` and `dN_IR` as `double`. The data type for states does not need to be defined at this stage and will be addressed later.
- ▶ `rbinom` is a built-in function used to generate random values from a binomial distribution. For additional built-in distributions in R, you can refer to this [Rmath.h](#) document.
- ▶ Remember to add a semicolon (`;`) after each line to ensure proper syntax.
- ▶ C snippets for the initializer and measurement model are:

## Specifying model components using C snippets IV

```
sir_rinit <- Csnippet("  
  S = nearbyint(Eta*N);  
  I = 1;  
  R = nearbyint((1-Eta)*N);  
  H = 0;  
")
```

```
sir_dmeas <- Csnippet("  
  lik = dnbinom_mu(reports,k,Rho*H,give_log);  
")
```

```
sir_rmeas <- Csnippet("  
  reports = rnbinom_mu(k,Rho*H);  
")
```

## Specifying model components using C snippets V

- ▶ No need to define the type for likelihood (`lik`) here, as it is already predefined.
- ▶ `nearbyint` is a built-in function used to find the closest integer to a given value.
- ▶ `reports` is the variable name specified in your dataset.



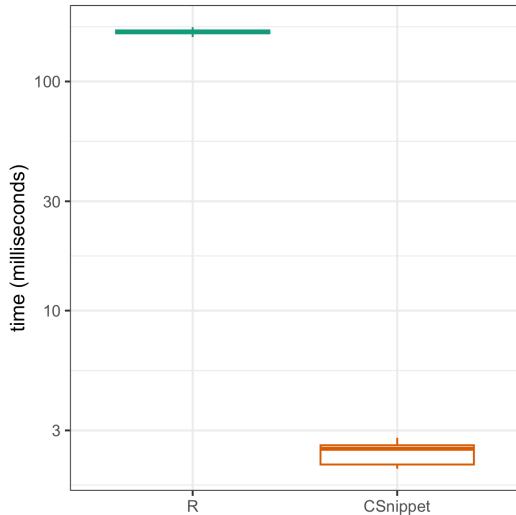
## Specifying model components using C snippets VI

- ▶ A call to `pomp` replaces the basic model components with these, much faster, implementations:

```
measSIR |>
  pomp(
    rprocess=euler(sir_stoch,delta.t=1/7),
    rinit=sir_rinit,
    rmeasure=sir_rmeas,
    dmeasure=sir_dmeas,
    accumvars="H",
    statenames=c("S","I","R","H"),
    paramnames=c("Beta","Gamma","N","Eta","Rho","k")
  ) -> measSIR_C
```

## Specifying model components using C snippets VII

- ▶ Note that, when using C snippets, one has to tell pomp which of the variables referenced in the C snippets are state variables and which are parameters. This is accomplished using the `statenames` and `paramnames` arguments.
- ▶ We can tell from the summary table that CSnippet is approximate 50 times faster than R.



## Guessing plausible parameter values I

- ▶ To check the code is working properly, we simulate. This requires us to assign parameters. A little thought will get us some ballpark estimates.
- ▶ Recall that  $\mathfrak{R}_0$  is the expected number of secondary infections resulting from one primary infection introduced into a fully susceptible population. For an SIR infection, one has that  $\mathfrak{R}_0 \approx \frac{L}{A}$ , where  $L$  is the lifespan of a host and  $A$  is the mean age of infection. Analysis of age-stratified serology data establish that the mean age of infection for measles during this period was around 4–5yr (Anderson and May 1991). Assuming a lifespan of 60–70yr, we have  $\mathfrak{R}_0 \approx 15$ .

## Guessing plausible parameter values II

- ▶ The basic theory of SIR epidemics gives the final-size equation,

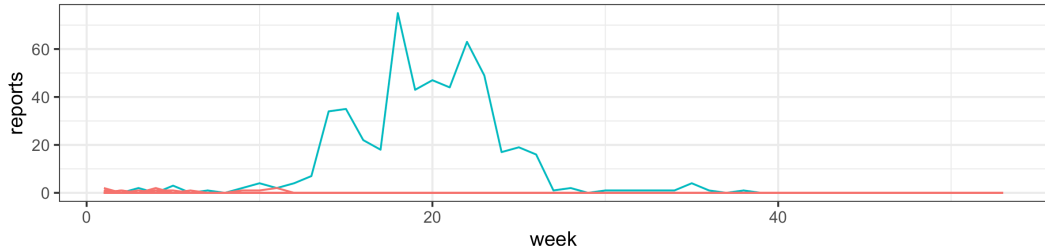
$$\mathfrak{R}_0 = -\frac{\log(1-f)}{f},$$

where  $f$  is the final size of the epidemic—the fraction of those susceptible at the beginning of the outbreak who ultimately become infected. For  $\mathfrak{R}_0 > 5$ , this equation predicts that  $f > 0.99$ .

- ▶ In the data, it looks like there were a total of 521 infections. Assuming 50% reporting, we have that  $S_0 \approx 1042$ , so that  $\eta = \frac{S_0}{N} \approx 0.027$ .
- ▶ If the infectious period is roughly 2 weeks, then  $1/\mu_{IR} \approx 2 \text{ wk}$  and  $\beta = \mu_{IR} \mathfrak{R}_0 \approx 7.5 \text{ wk}^{-1}$ .

## Guessing plausible parameter values III

```
measSIR |>  
  simulate(params=c(Beta=7.5,Gamma=0.5,Rho=0.5,k=10, Eta=0.03,N=38000),  
    nsim=20,format="data.frame",include.data=TRUE) |>  
  ggplot(aes(x=week,y=reports,group=.id,color=.id=="data")) +  
  geom_line() + guides(color="none")
```



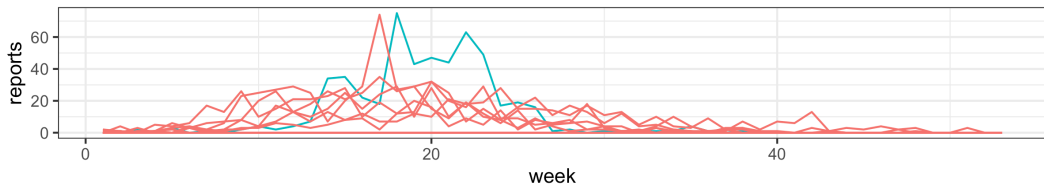
## Exercise I: Explore the SIR model

Fiddle with the parameters to see if you can't find a model for which the data are a more plausible realization.

## Worked solutions I: Explore the SIR model I

In the simulated outbreaks, the overall incidence is much too low, and the outbreak dies out immediately. We might try increasing the force of infection:

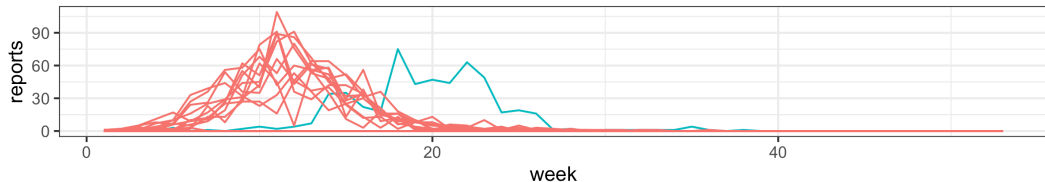
```
measSIR |>  
  simulate(params=c(Beta=25, Gamma=0.5, Rho=0.5, k=10, Eta=0.03, N=38000),  
    nsim=20, format="data.frame", include.data=TRUE) |>  
  ggplot(aes(x=week, y=reports, group=.id, color=.id=="data")) +  
  geom_line() + guides(color="none")
```



## Worked solutions I: Explore the SIR model II

Taking it farther...

```
measSIR |>  
  simulate(params=c(Beta=40,Gamma=0.5,Rho=0.5,k=10,Eta=0.03,N=38000),  
    nsim=20,format="data.frame",include.data=TRUE) |>  
  ggplot(aes(x=week,y=reports,group=.id,color=.id=="data")) +  
  geom_line() + guides(color="none")
```

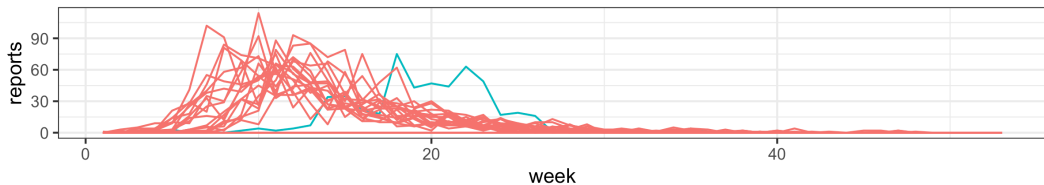




## Worked solutions I: Explore the SIR model III

While this increases the overall incidence, the epidemic is now peaking too quickly. To counteract this, we might try reducing the recovery rate.

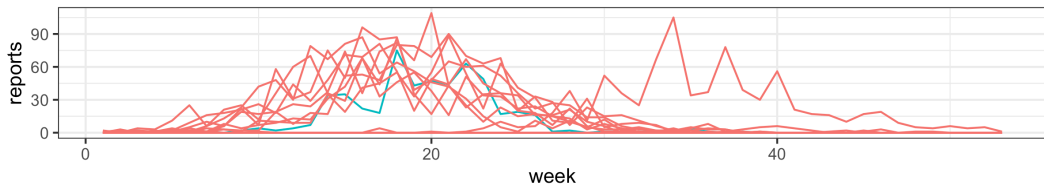
```
measSIR |>  
  simulate(params=c(Beta=40,Gamma=0.2,Rho=0.5,k=10,Eta=0.03,N=38000),  
    nsim=20,format="data.frame",include.data=TRUE) |>  
  ggplot(aes(x=week,y=reports,group=.id,color=.id=="data"))+  
  geom_line() + guides(color="none")
```



## Worked solutions I: Explore the SIR model IV

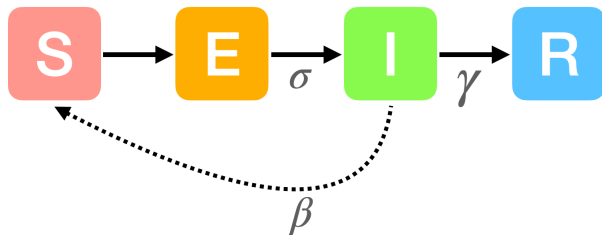
Additionally, we might have a look at the effects of changing the initial susceptible fraction,  $\eta$ :

```
measSIR |>  
  simulate(params=c(Beta=15, Gamma=0.5, Rho=0.5, k=10, Eta=0.06, N=38000),  
    nsim=20, format="data.frame", include.data=TRUE) |>  
  ggplot(aes(x=week, y=reports, group=.id, color=.id=="data")) +  
  geom_line() + guides(color="none")
```



## Exercise II: Extend the SIR model to SEIR model

Below is a diagram of the so-called SEIR model. This differs from the SIR model in that infected individuals must pass a period of latency before becoming infectious.



Modify the codes above to construct a pomp object containing the Consett measles data and an SEIR model. Perform simulations as above and adjust parameters to get a sense of whether improvement is possible by including a latent period.

## Worked solutions II: The SEIR model I

The existing code may be modified as follows:

```
seir_stoch <- Csnippet("  
  double dN_SE = rbinom(S,1-exp(-Beta*I/N*dt));  
  double dN_EI = rbinom(E,1-exp(-Sigma*dt));  
  double dN_IR = rbinom(I,1-exp(-Gamma*dt));  
  S -= dN_SE;  
  E += dN_SE - dN_EI;  
  I += dN_EI - dN_IR;  
  R += dN_IR;  
  H += dN_IR;  
")
```

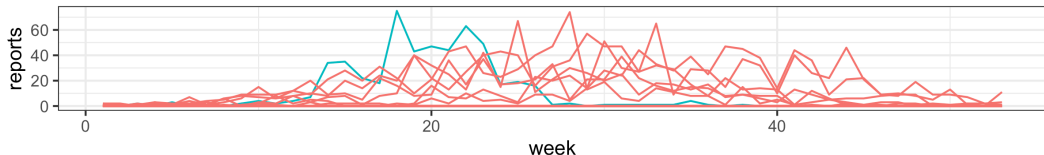
## Worked solutions II: The SEIR model II

```
seir_init <- Csnippet("  
  S = nearbyint(Eta*N);  
  E = 0; I = 1;  
  R = nearbyint((1-Eta)*N);  
  H = 0;  
")  
  
measSIR |>  
  pomp(  
    rprocess=euler(seir_stoch,delta.t=1/7),  
    rinit=seir_init,  
    paramnames=c("N","Beta","Sigma","Gamma","Rho","Eta","k"),  
    statenames=c("S","E","I","R","H")  
  ) -> measSEIR
```

## Worked solutions II: The SEIR model III

Using the rough estimate that the latent period in measles is 8–10da, we take  $\sigma \sim 0.8\text{wk}^{-1}$  and  $\gamma \sim 1.3\text{wk}^{-1}$  (roughly the same generation time as before).

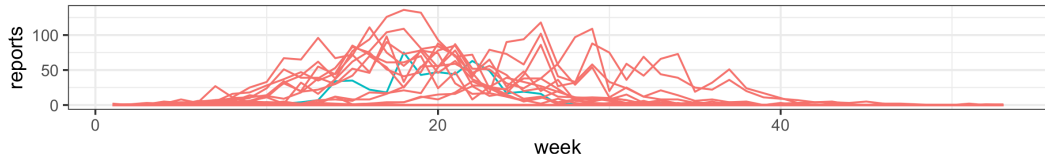
```
measSEIR |>
  simulate(params=c(Beta=30,Sigma=0.8,Gamma=1.3,
                    Rho=0.5,k=10,Eta=0.06,N=38000),
    nsim=20,format="data.frame",include.data=TRUE) |>
  ggplot(aes(x=week,y=reports,group=.id,color=.id=="data")) +
  geom_line() + guides(color="none")
```



## Worked solutions II: The SEIR model IV

Again one can increase the force of infection:

```
measSEIR |>
  simulate(params=c(Beta=40,Sigma=0.8,Gamma=1.3,
                    Rho=0.5,k=10,Eta=0.06,N=38000),
    nsim=20,format="data.frame",include.data=TRUE) |>
  ggplot(aes(x=week,y=reports,group=.id,color=.id=="data")) +
  geom_line() + guides(color="none")
```




## References

- Anderson, R. M., and R. M. May. 1991. *Infectious Diseases of Humans*. Oxford: Oxford University Press.
- Bretó, Carles, Daihai He, Edward L. Ionides, and Aaron A. King. 2009. "Time Series Analysis via Mechanistic Models." *Ann Appl Stat* 3 (1): 319–48.  
<https://doi.org/10.1214/08-AOAS201>.



## License, acknowledgments, and links

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- ▶ Produced with R version 4.3.2 and pomp version 5.10.
- ▶ Compiled on 2024-07-24.

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