### Case study: Spanish Flu in London, Birmingham, and Liverpool

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# **Objectives**

- 1. To explore the use of POMP models in the context of meta-populations.
- 2. To illustrate the use of POMP models in a more complex setting.
- 3. To demonstrate the use of diagnostic probes for model criticism.
- 4. To provide an example that can be modified to apply similar approaches to other outbreaks of emerging infectious diseases.

# Emerging infectious disease pandemic I

- ▶ The 1918 Spanish flu pandemic was one of the deadliest pandemics in human history.
- ▶ The pandemic lasted from January 1918 to December 1920, spreading to nearly every part of the world.
- $\blacktriangleright$  The pandemic was caused by the H1N1 influenza A virus.
- ▶ The pandemic was responsible for the deaths of an estimated 50 million people worldwide.

# Emerging infectious disease pandemic II

Key questions included:

- 1. How fast will the pandemic unfold?
- 2. What are the common and differentiated characteristics of the pandemic in different cities?
- 3. How people will correspond to the pandemic?

#### Weekly reported data I

dat <- read\_csv("1918flu\_3cities\_1wave.csv") head(dat)



# Weekly reported data II



### SIR model with behavioral response I

- ▶ Social distancing is a common public health intervention.
- $\triangleright$  We model the effect of social distancing as a reduction in the transmission rate.
- $\blacktriangleright$  The SIR model with behavioral response is given by:



#### SIR model with behavioral response II

$$
\frac{dS}{dt} = -\beta(t) S \frac{I}{N}
$$
  
\n
$$
\frac{dI}{dt} = \beta(t) S \frac{I}{N} - \gamma I
$$
  
\n
$$
\frac{dR}{dt} = (1 - \phi) \gamma I
$$
  
\n
$$
\frac{dD}{dt} = \phi \gamma I - g D
$$
  
\n
$$
\frac{dM}{dt} = g D
$$
  
\n
$$
\frac{dP}{dt} = g D - \lambda P
$$

- $\triangleright$  S, I, R: the susceptible, infectious, and recovered populations
- $\triangleright$  D: those who have lost of infectiousness and are progressing to death due to influenza or/and pneumonia
- $\blacktriangleright M$ : those who have died
- $\blacktriangleright$  P: general public's risk perception based on recent influenza deaths
- $\triangleright$   $N: N = S + I + R + D + M$ , the constant total population size

### SIR model with behavioral response III

The time-dependent transmission rate  $\beta(t)$  consists of two components:

$$
\beta(t) = \beta_0 \cdot \left(1 - \frac{P(t)}{N}\right)^{\kappa} \tag{1}
$$

 $\blacktriangleright$   $\beta_0$ : the baseline transmission rate ▶  $\left(1-\frac{P(t)}{N}\right)^{\kappa}$ : represent the effect of reactive social distancing on transmission rate based on the public's risk perception  $P(t)$ 

The number of reported cases  $C$  follows a Negative Binomial distribution, in London, Birmingham, and Liverpool, respectively, with ratio  $\rho$  and over-dispersion parameter k, given the cumulative number of cases  $H$ :

 $C \sim$  NegBin $(\rho H, k)$ 

# SIR model with behavioral response IV

- ▶ Fixed parameters:
	- ▶ The total population sizes for London, Birmingham, and Liverpool are fixed at 4484523, 919444, and 802940, respectively
	- $\blacktriangleright$   $\gamma^{-1}$ : the average infectious period, fixed at 4 days
	- ▶  $g^{-1}$ : the mean time from loss-of-infectiousness to death, fixed at 8 days
- $\blacktriangleright$  Parameters to be estimated, initial conditions:
	- $\blacktriangleright \eta_L, \eta_R, \eta_L$ : the initial fraction of susceptible individuals for London, Birmingham, and Liverpool, respectively
	- $\blacktriangleright \psi_L, \psi_B, \psi_L$ : the initial fraction of infectious individuals for London, Birmingham, and Liverpool, respectively
- ▶ Parameters to be estimated, common features:
	- $\blacktriangleright$   $\beta_0$ : the baseline transmission rate
	- $\triangleright$   $\kappa$ : the exponent of the social distancing effect
	- $\blacktriangleright$   $\lambda$ : the rate at which the public's risk perception decays

The implementation in pomp: the state process for meta-populations

```
sir meta <- Csnippet("
  double *S = \&S1, *I = \&I1, *R = \&R1, *D = \&D1, *M = \&M1;double *P = \&P1, *H = \&H1; int N[3] = \{N1, N2, N3\};
  double Beta, dN_SI, dN_IRD, dN_IR, dN_ID, dN_DM, dN_P;
  for (int i = 0; i < 3; i++) {
   Beta = beta0*pow(1-P[i]/N[i], kappa);
   dN SI = rbinom(S[i],1-exp(-Beta*I[i]/N[i]*dt));dN IRD = rbinom(I[i], 1-exp(-gamma*dt));
    dN IR = nearbyint((1-phi)*dN IRD); dN ID = nearbyint(phi*dN IRD);
    dN DM = rbinom(D[i], 1-exp(-g*dt));
   dN P = rbinom(P[i], 1-exp(-lambda*dt));
   S[i] -= dN_SI; I[i] += dN_SI - dN_IRD; R[i] += dN_IR;
   D[i] += dN_ID - dN_DM; M[i] += dN_DM; P[i] += dN_DM - dN_P;
   H[i] += dN IRD; }
")
```
### The implementation in pomp: the initial conditions

```
sir meta rinit <- Csnippet("
  double *S = \&S1, *T = \&I1, |R = \&R1, *D = \&D1, *M = \&M1;
  double *P = \&P1, *H = \&H1; int N[3] = \{N1, N2, N3\};
  double eta[3] = \{eta1, eta2, eta3\};
  double psi[3] = \{psi1,psi2,psi3\};for (int i = 0; i < 3; i++) {
    S[i] = nearbyint(N[i]*eta[i]);I[i] = nearbyint(N[i]*psi[i]);
   R[i] = nearbyint(N[i]*(1-eta[i]-psi[i]));
   D[i] = M[i] = P[i] = H[i] = 0;
 }
")
```
### The implementation in pomp: the measurements

```
sir meta dmeas <- Csnippet("
 double lik1, lik2, lik3;
 lik1 = (ISMA(London)) ? 0 : dhbinommu(London,rho*H1,k,1);lik2 = (ISMA(Birmingham)) ? 0 : dnbinom mu(Birmingham,rho*H2,k,1);
 lik3 = (ISMA(Liverpool)) ? 0 : dnbinom mu(Liverpool,rho*H3,k,1);
 lik = lik1 + lik2 + lik3;lik = (give log) ? lik : exp(lik);
")
sir meta rmeas <- Csnippet("
 London = rnbinom mu(k,rho*H1);Birmingham = rnbinom mu(k,rho*H2);
 Liverpool = rnbinom_mu(k,rho*H3);")
```
The implementation in pomp: build the model I

```
dat |> select(-date) |>
  pomp(
    times = "week", t0 = 0.
    rprocess=euler(sir_meta,delta.t=1/7),
    rinit=sir_meta_rinit, rmeasure=sir_meta_rmeas,
    dmeasure=sir meta dmeas, accumvars = sprintf("H%d",1:3),
    statenames=c(sprintf("S%d",1:3),sprintf("I%d",1:3),
      sprintf("R''_0d'', 1:3), sprintf("D''_0d'', 1:3), sprintf("M''_0d'', 1:3),
      sprintf("P%d",1:3),sprintf("H%d",1:3)),
    paramnames=c(
      "beta0","kappa","gamma","phi","g","lambda","rho","k",
      sprintf("N%d",1:3),sprintf("eta%d",1:3),sprintf("psi%d",1:3))
  ) -> pomp_meta
```
# The implementation in pomp: build the model II

```
params \leq -c (beta0 = 5, kappa = 5, gamma = 1/4, phi = 0.0119, g = 1/8,
 lambda = 1/2, rho = 0.05, k = 10, N1 = 4484523, N2 = 919444,
  N3 = 802940, eta1 = 0.2, eta2 = 0.2, eta3 = 0.2,
  psi1 = 0.0005, psi2 = 0.0005, psi3 = 0.0005)
pomp meta |>simulate(
   params = params, nsim=20,format="data.frame",include.data=TRUE
  ) |>select(week,.id,London,Birmingham,Liverpool) |>
  reshape2::melt(id.vars = c("week", ".id")) |>
 ggplot(aes(x=week, y=value, group=.id, color=.id=="data")) +
  geom_line() + facet_wrap(variable \sim .) +
  theme minimal() + guides(color="none")
```
# The implementation in pomp: build the model III



#### License, acknowledgments, and links

- ▶ This lesson is prepared for the Simulation-based Inference for Epidemiological Dynamics module at the Summer Institute in Statistics and Modeling in Infectious Diseases, SISMID.
- ▶ The materials build on previous versions of this course and related courses.
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- ▶ Produced with R version 4.4.0 and pomp version 5.9.
- ▶ Compiled on 2024-07-24.

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