UNIT 8A: Infectious disease

1 Introduction

Infectious disease

- Extremely common
- Huge impacts on ecological interactions
- A form of exploitation, but doesn't fit well into our previous modeling framework
  - How many people are there?
  - How many influenza viruses are there?
  - How do they find each other?

Disease agents

- Poll: Name an infectious agent that causes disease in humans.
- Disease agents vary tremendously:
  - Most **viruses** have just a handful of genes that allow them to hijack a cell and get it to make virus copies
    - *
  - **Bacteria** are independent, free-living cells with hundreds or thousands of chemical pathways
    - *
  - **Eukaryotic** pathogens are nucleated cells who are more closely related to you than they are to bacteria
    - *

Microparasites

- For infections with small pathogens (viruses and bacteria), we don't attempt to count pathogens, but instead divide disease into stages
  - Latently infected
  - Productively infected
  - Recovered

Microparasite models

- We model microparasites by counting the number of hosts in various **states**:
  - **Susceptible** individuals can become infected
  - **Infectious** individuals are infected and can infect others
  - **Resistant** individuals are not infected and cannot become infected
- More complicated models might include other states, such as latently infected hosts who are infected with the pathogen but cannot yet infect others

Models as tools

- Models are the tools that we use to connect scales:
  - individuals to populations
  - single actions to trends through time

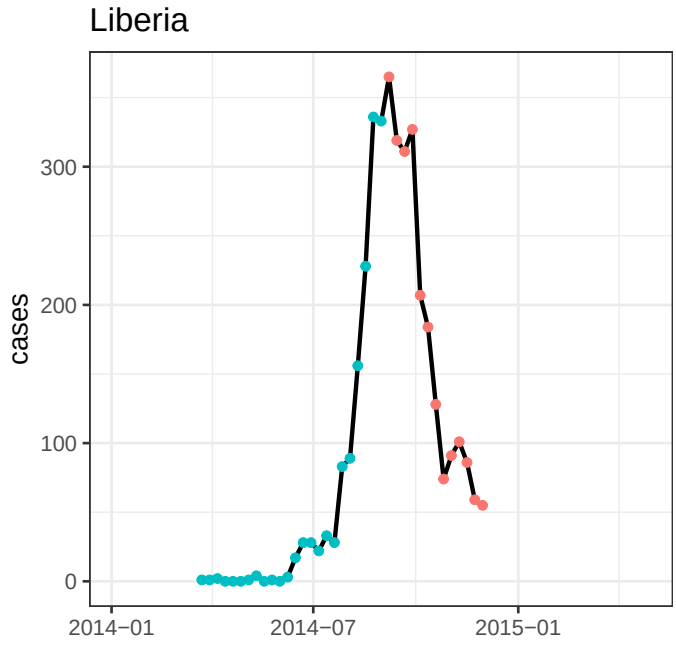
2 Rate of spread

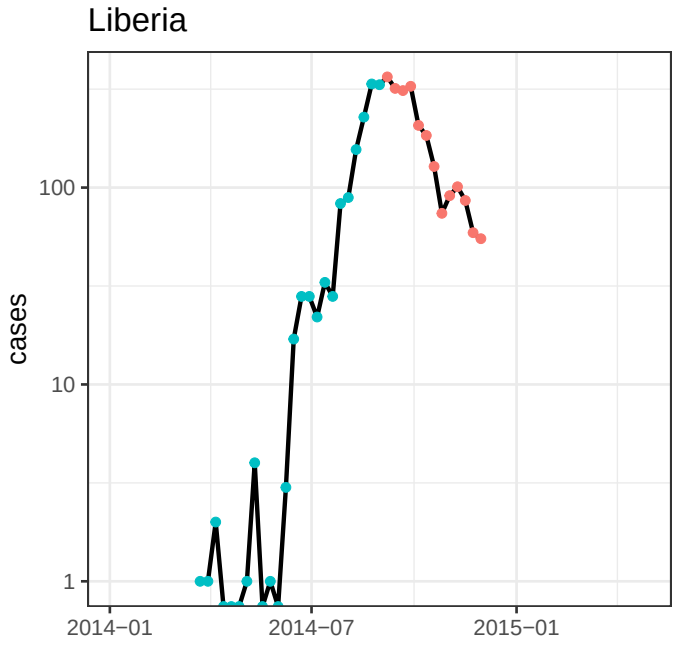
- Poll: For many diseases, especially new diseases, we can *observe* and *estimate* r.
  -
    - *
    - *
- Poll: Want to know what factors contribute to that, and how it relates to $\mathcal{R}$.
  -
  -

Basic reproductive number

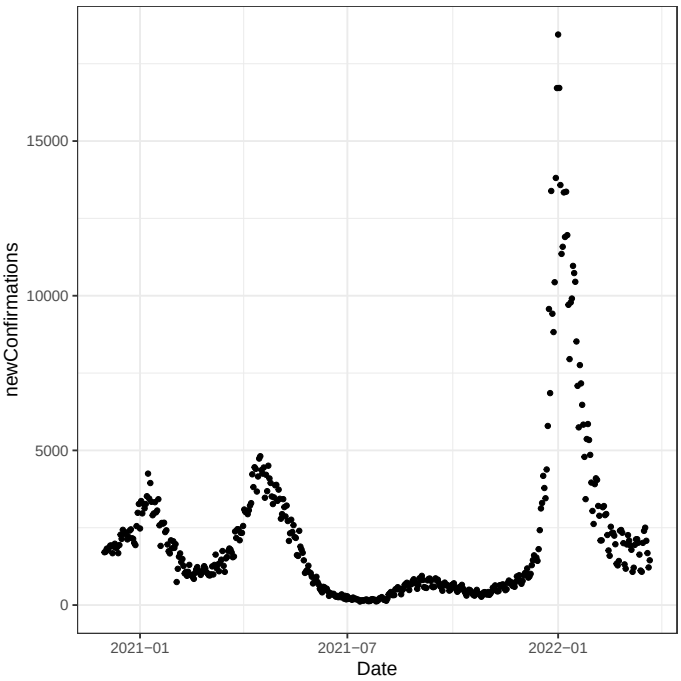
- People in the disease field love to talk specifically about $\mathcal{R}_0$
- But they don't always mean the same thing when they say $\mathcal{R}_0$:
  - Actual value of $\mathcal{R}$ before an epidemic
  - Hypothetical value assuming no immunity
  - Hypothetical value assuming no immunity and no control efforts whatsoever
- Often easier to talk simply about $\mathcal{R}$.

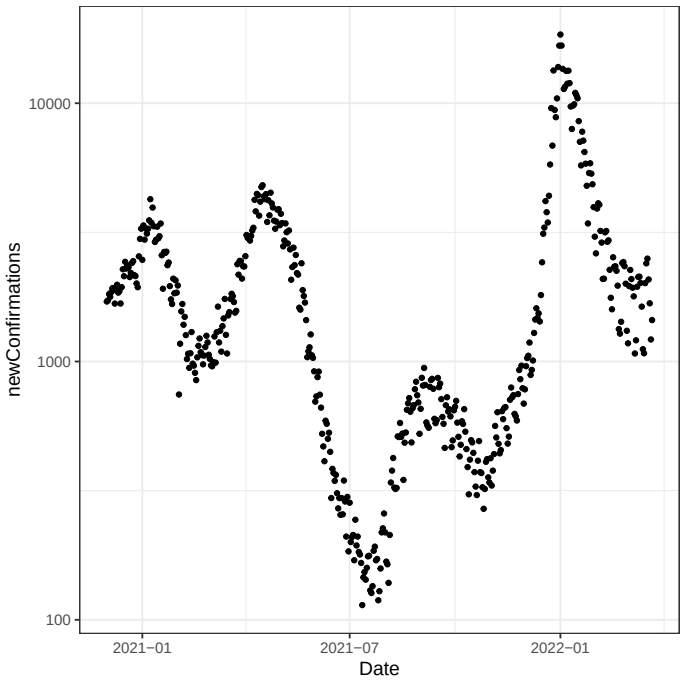
Example: the West African Ebola epidemic





COVID in Ontario





Scales

- Which scale should we look at?

  –

  –

Population biology

- What quantities do we want to look at?
  -
  -
    -
  -

Instantaneous rate of growth r

- What are the components?
  -
    -
    -
  -
    -
    -
- How do you think we estimate?
  -
    -
  -
    - Individuals $\leftrightarrow$ Populations

Finite rate of growth λ

- Why do we want this?
  -
  -
- How do we calculate it?
  -
  -

Example

- $r \approx 0.14/$ day for early COVID spread
- What is λ?
  - At a time scale of a day?
  - At a time scale of a week?

Reproductive number $\mathcal{R}$

- What is it?
  - –
- Why do we want this?
  - –
- How do we calculate it?
  - –

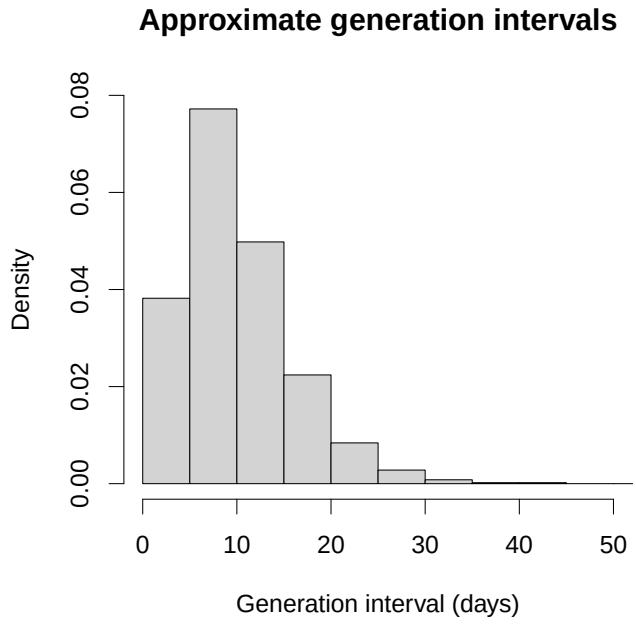
Example

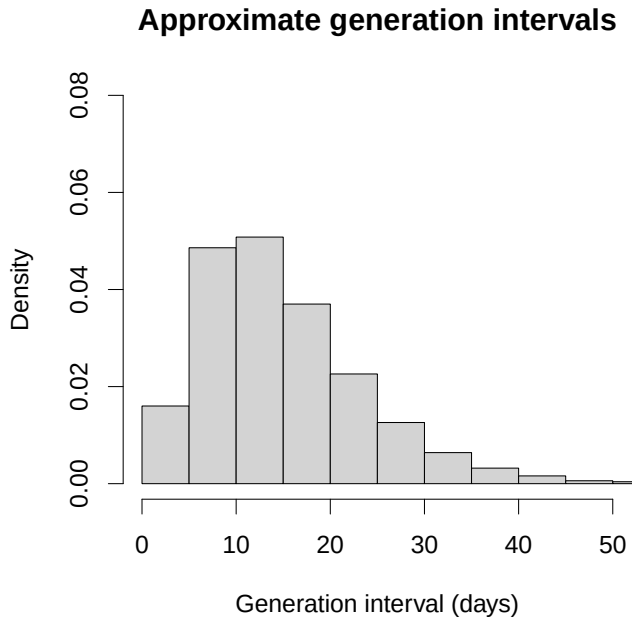
- $r \approx 0.14/\,\text{day}$
- What is our estimate of $\mathcal{R}$?
  - When average length of infection $L = 5\,\text{day}$?
  - When average length of infection $L = 10\,\text{day}$?

Generation intervals

- Researchers try to estimate the *proportion* of transmission that happens for different **ages of infection**
- How long from the time you are *infected* to the time you *infect someone else*?
- Analogous to a life table
- The effective generation time $\hat{G}$ has units of time

Generation intervals





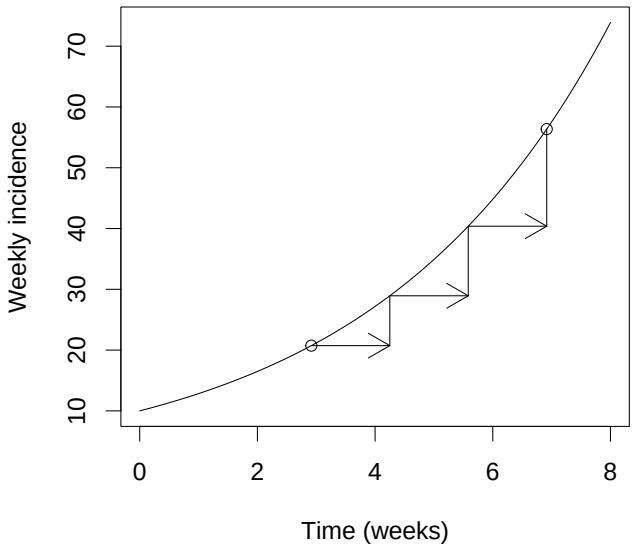
Speed and risk

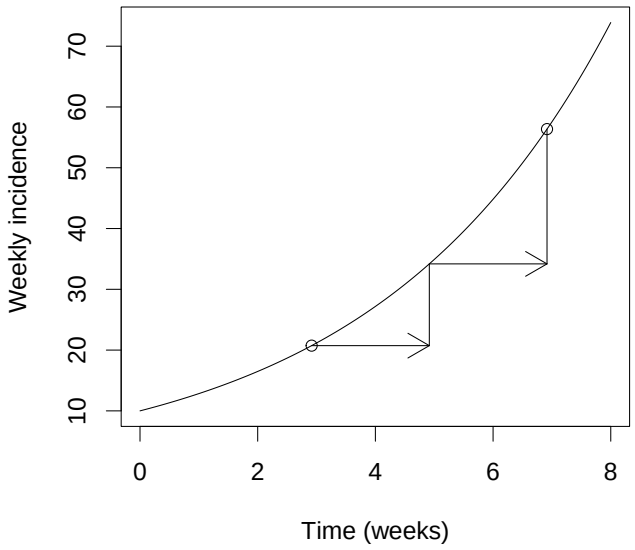
- Which is more dangerous, a fast disease, or a slow disease?
  - How are we measuring speed?
  - How are we measuring danger?
  - *What do we already know?*

Generation time and risk

- If we know $\mathcal{R}$, what does the generation time tell us about r?
  -
- If we know r, what does the generation time tell us about $\mathcal{R}$?
  -
- $\mathcal{R} = \exp(r\hat{G})$

Generation time and risk





Generation time and risk

- $\mathcal{R} = \exp(r\hat{G})$
- An intuitive view:
  - Epidemic speed = Generation strength $\times$ Generation speed
- If we know generation speed, then a faster epidemic speed means:

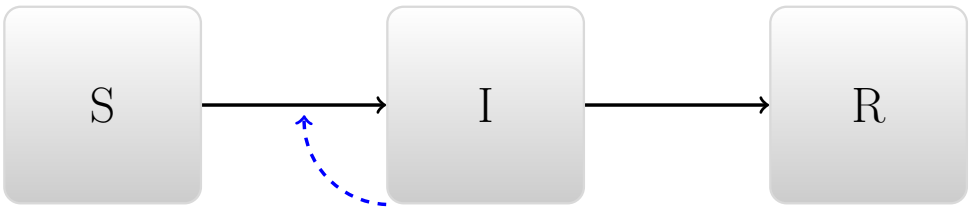
–

- If we know epidemic speed, a faster generation speed means
  –

3 Single-epidemic model

- Susceptible → Infectious → Recovered
- We also use N to mean the total population

Transition rates



- What factors govern movement through the boxes?
  - People get better independently
  - People get infected by infectious people

Conceptual modeling

- Poll: What happens in the long term if we introduce an infectious individual?
  –
  –
  –
    *

Interpreting

- Why might there not be an epidemic?
  -
    - *
  -
- Why doesn't everyone get infected?
  -

Implementing the model

- The simplest way to implement this conceptual model is with differential equations:
  - $$\frac{dS}{dt} = -\beta \frac{SI}{N}$$
  - $$\frac{dI}{dt} = \beta \frac{SI}{N} - \gamma I$$
  - $$\frac{dR}{dt} = \gamma I$$
  - $$N = S + I + R$$

Quantities

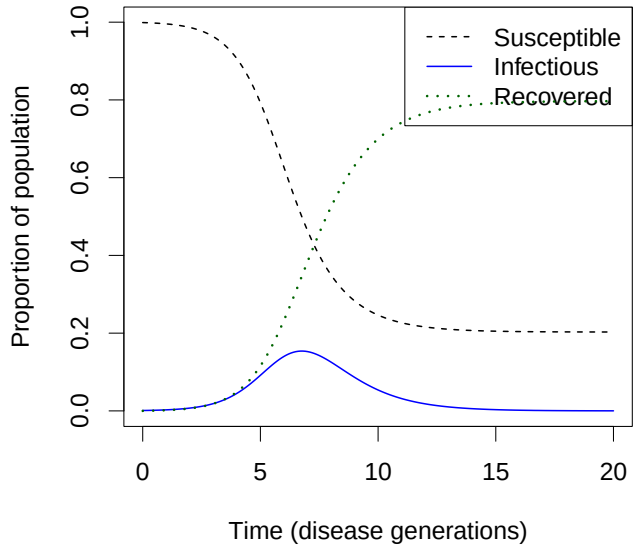
State variables

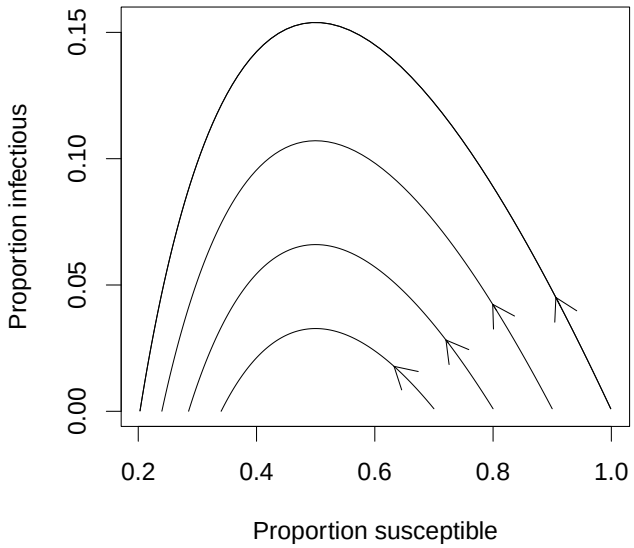
- S, I, R, N: [people] or [people/ha]

Parameters

- Susceptible people have **potentially effective** contacts at rate β (units [1/time])
  - These are contacts that would lead to infection if the person contacted is infectious
  - Total infection rate is $\beta I/N$, because I/N is the proportion of the population infectious
- Infectious people recover at *per capita* rate γ (units [1/time])
  - Total recovery rate is γI
  - Mean time infectious is $D = 1/\gamma$ (units [time])

Simulating the model





Basic reproductive number

- Poll: What *unitless* parameter can you make from the model above?
  - –
  - –
    - *
  - –

Basic reproductive number implications

- Poll: What happens early in the epidemic if $\mathcal{R}_0 > 1$?
  - –
- What happens early in the epidemic if $\mathcal{R}_0 < 1$?
  - –

Effective reproductive number

- The effective reproductive number gives the number of new infections per infectious individual in a partially susceptible population:
  - –
- Is the disease increasing or decreasing?
  - –

  -

- Why doesn't everyone get infected?
  -
  -

3.1 Epidemic size

- In this model, the epidemic always burns out
  - No source of new susceptibles
- Epidemic size is determined by:
  -
  -
    *
    *
  -
    *

Overshoot

- Why does more susceptibles at the beginning mean fewer susceptibles at the end?
  -
  -
  -
  -
  -
  -

Ebola example

- In September, the US CDC predicted "as many as" 1.5 million Ebola cases in Liberia by January
- In fact, their model predicted many *more* cases than that by April
- What happened?

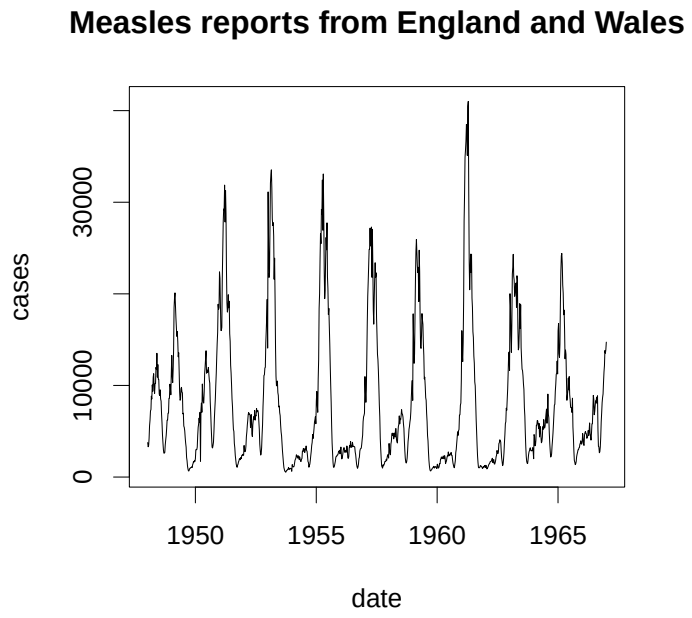
What limits epidemics?

- Poll: What limits epidemics in our simple models?
  –
- Poll: What else limits epidemics in real life?
  –
  –
  –

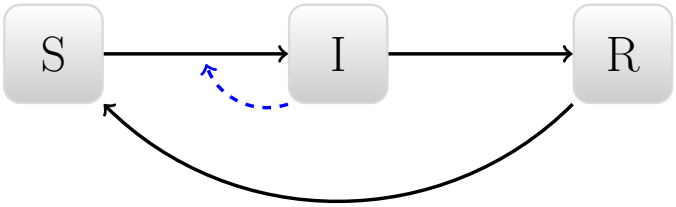
4 Recurrent epidemic models

- Poll: If epidemics tend to burn out, why do we often see repeated epidemics?
  –
  –

Recurrent epidemics

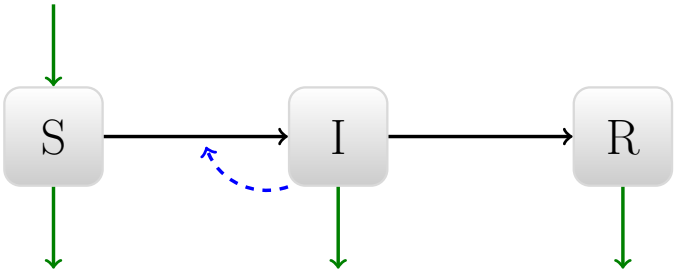


Closing the circle



-

Closing the circle



-
  -

Births and deaths

- $$\frac{dS}{dt} = bN - \beta\frac{SI}{N} - dS$$
- $$\frac{dI}{dt} = \beta\frac{SI}{N} - \gamma I - dI$$

- $$\frac{dR}{dt} = \gamma I - dR$$
- We often assume $b = d$
  - $\implies$ population is constant

4.1 Dynamics

Equilibrium

- At equilibrium, we know that $\mathcal{R}_{\text{eff}} = 1$
  - One case per case
  - Number of susceptibles at equilibrium determined by the number required to keep infection in balance
    - $S/N = 1/\mathcal{R}_0$
- What does this remind you of?
  -
- Number of infectious individuals determined by number required to keep susceptibles in balance.
- As susceptibles go up, what happens?
  - Per capita replenishment goes down
  - Infections required goes down

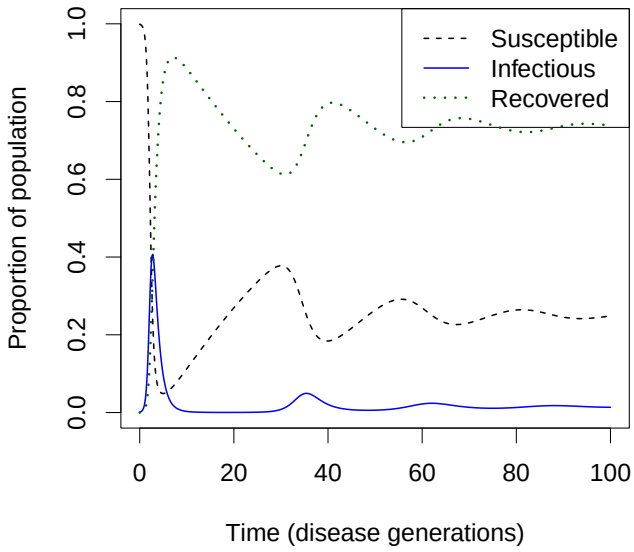
Reciprocal control

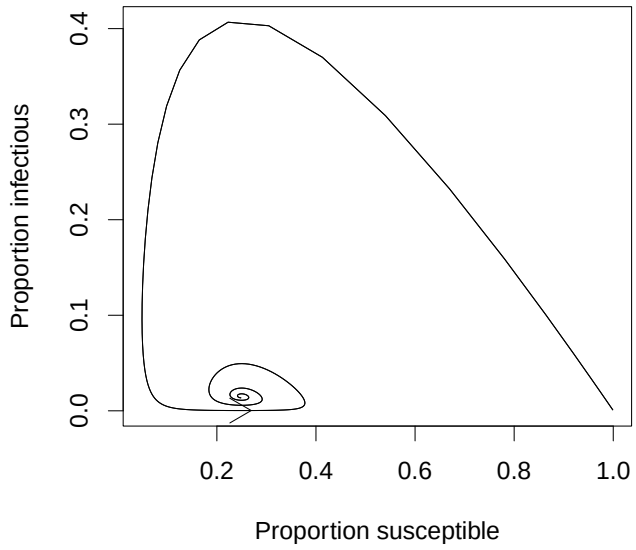
- What happens to *equilibrium* if we protect susceptibles (move them to R class)?
  -
  -
  -
  -
- What else could happen?
  -
  -
    *

Reciprocal control

- Poll: What happens if we remove infectious individuals at a constant rate (find them and cure them or isolate them)?
  -
  -
    -
  -

Tendency to oscillate





Tendency to oscillate

- "Closed-loop" SIR models (ie., with births or loss of immunity):
  - Tend to oscillate
  - Oscillations tend to be damped
    - System reaches an **endemic** equilibrium – disease persists

Source of oscillations

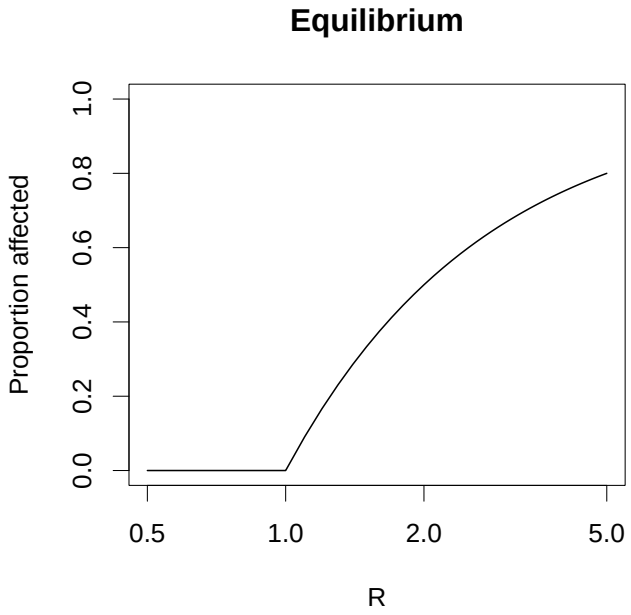
- Similar to predator-prey systems
- What happens if we start with too many susceptibles?
  -
  -
  -
  -

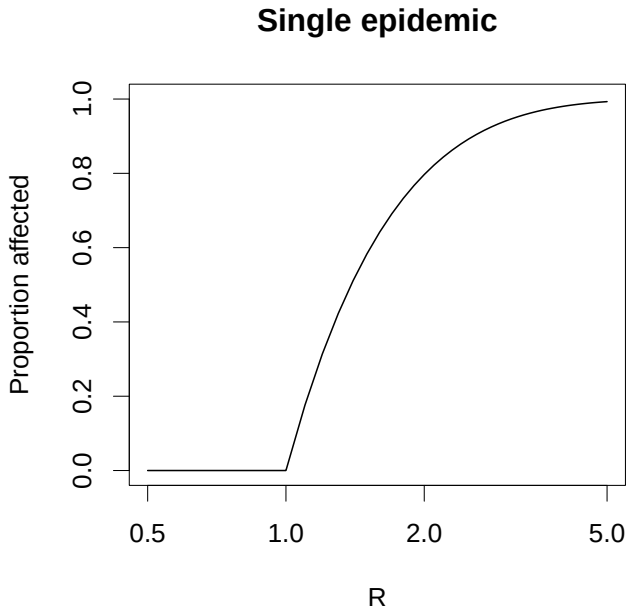
Persistent oscillations

- Poll: If oscillations tend to be damped in simple models, why do they persist in real life?
  -
    - *
    - *
  -
  -
  -
    - *

5 Reproductive numbers and risk

- At equilibrium, the proportion of people who are susceptible to disease should be approximately $S/N = 1/\mathcal{R}_0$
- Proportion "affected" (infectious or immune) should be approximately $V/N = 1-1/\mathcal{R}_0$
- If you have a single, fast epidemic, the size is also predicted by $\mathcal{R}_0$.





Examples

- Ronald Ross predicted 100 years ago that reducing mosquito densities by a factor of 5 or so would *eliminate* malaria
- Gradual disappearance of polio, typhoid, etc., without risk factors going to zero
- Eradication of smallpox!

Threshold for elimination

- What proportion of the population should be vaccinated to eliminate a disease?
  –

Examples:

- Polio has an $\mathcal{R}_0$ of about 5.
- Poll: What proportion of the population should be vaccinated to eliminate polio?
  –
- Measles has an $\mathcal{R}_0$ of about 20. What proportion of the population should be vaccinated to eliminate measles?
  –
- If gonorrhea has an $\mathcal{R}_0$ of about 2, what proportion of unprotected sexual encounters should be protected to eliminate gonorrhea?
  –

Persistence of infectious disease

- Why have infectious diseases persisted?
  - The pathogens *evolve*
  - Human populations are **heterogeneous**
    - People differ in: nutrition, exposure, access to care
  - Information and misinformation
    - Vaccine scares, trust in health care in general

Heterogeneity and persistence

- Heterogeneity *increases* $\mathcal{R}_0$
  - When disease is rare, it is concentrated in the most vulnerable populations
    - Cases per case is high
    - Elimination is harder
- Marginal populations
  - Heterogeneity could make it easier to concentrate on the most vulnerable populations and eliminate disease
  - Humans rarely do this, however: the populations that need the most support typically have the least access