

Population heterogeneity

- What are the effects of population heterogeneity (differences between individuals) on disease spread?

● Population heterogeneity

The implications of homogeneity

The effects of heterogeneity

Population heterogeneity

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The effects of heterogeneity

- What are the effects of population heterogeneity (differences between individuals) on disease spread?
- What are the effects of homogeneity?

- Population heterogeneity

The implications of homogeneity

- Equilibrium analysis
- The endemic curve

The effects of heterogeneity

The implications of homogeneity

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The effects of heterogeneity

- What conclusions can we reach if we assume that people are effectively the same for the purposes of transmitting a particular disease?

The basic reproductive number

- $\mathcal{R}_0$ is the number of people who would be infected by an infectious individual in a susceptible population.

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The basic reproductive number

- $\mathcal{R}_0$ is the number of people who would be infected by an infectious individual in a susceptible population.
- $\mathcal{R}_0 = pcD = \beta D = \beta/\gamma$.

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- $\mathcal{R}_0$ is the number of people who would be infected by an infectious individual in a susceptible population.
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 - ◆ p : Probability of transmission

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 - ◆ p : Probability of transmission
 - ◆ c : Contact Rate
 - ◆ D : Average duration of infection

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- $\mathcal{R}_0$ is the number of people who would be infected by an infectious individual in a susceptible population.
- $\mathcal{R}_0 = pcD = \beta D = \beta/\gamma$.
 - ◆ p : Probability of transmission
 - ◆ c : Contact Rate
 - ◆ D : Average duration of infection
- A disease can invade a population $\iff \mathcal{R}_0 > 1$.

Equilibrium analysis

- More generally, the number of new cases per case is

$$\mathcal{R} = pcD \frac{S}{N} = \beta D \frac{S}{N} = \mathcal{R}_0 \frac{S}{N}.$$

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- At equilibrium: $\mathcal{R} = \mathcal{R}_0 \frac{S}{N} = 1.$

Equilibrium analysis

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- Thus: $\frac{S}{N} = 1/R_0.$

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- At equilibrium: $\mathcal{R} = \mathcal{R}_0 \frac{S}{N} = 1$.

- Thus: $\frac{S}{N} = 1/R_0$.

- Proportion ‘affected’ is $V = 1 - S/N = 1 - 1/R_0$.

The endemic curve

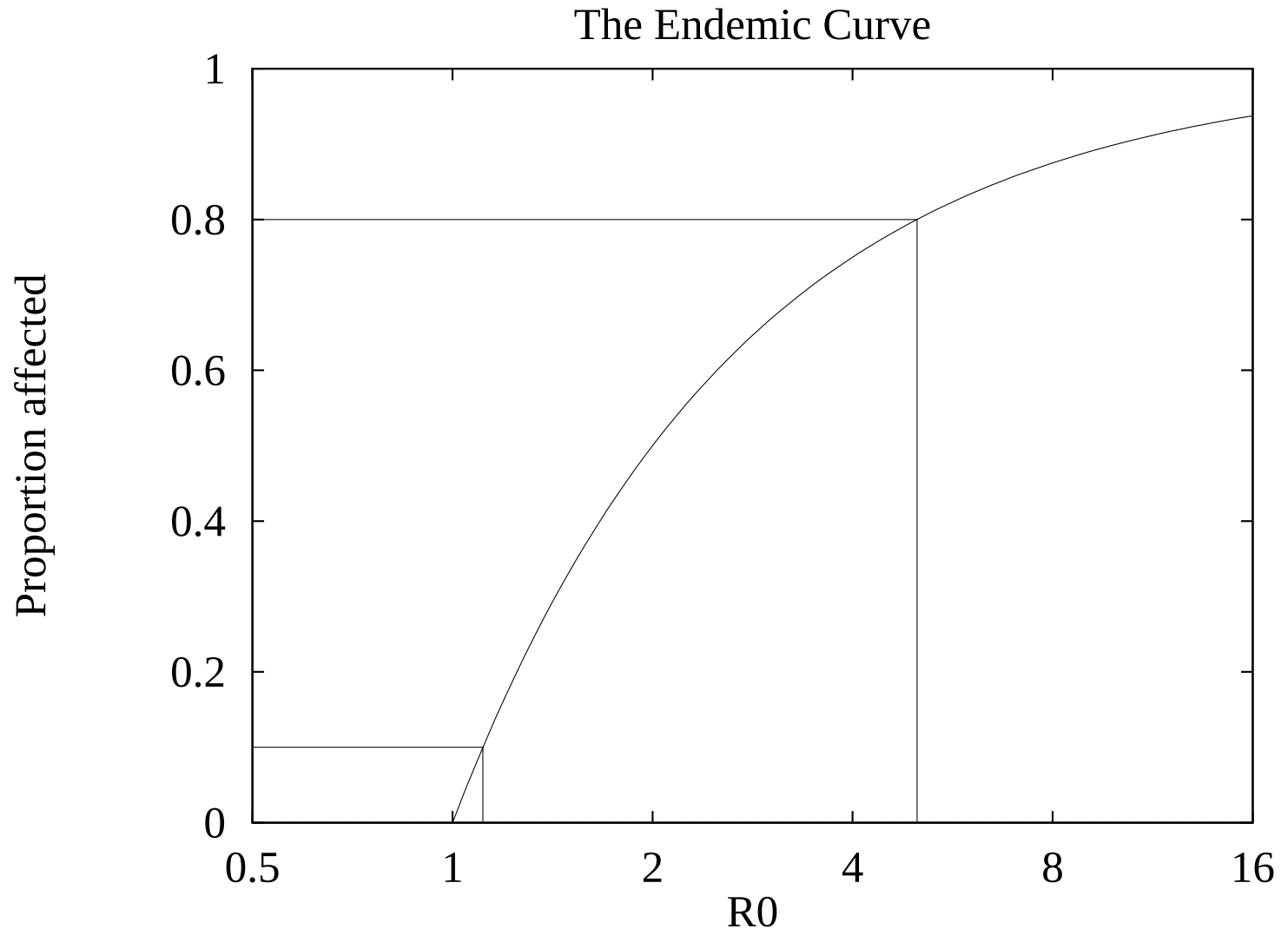
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The effects of heterogeneity



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- Disease Heterogeneity in TB
- Heterogeneity in other diseases
- Next-generation framework
- Heterogeneity and $\mathcal{R}_0$
- Heterogeneity and equilibrium

The effects of heterogeneity

Beyond the homogeneous model

■ Heterogeneity among hosts

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- Demographic heterogeneity (individuality)

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 - ◆ “waiting” distributions

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- Heterogeneity among hosts
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- Demographic heterogeneity (individuality)
- Temporal heterogeneity
 - ◆ “waiting” distributions
 - ◆ seasonality

Heterogeneity among hosts

- Differences among people are pervasive, large and often correlated

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- We break down the transmission probability into a focus and a victim' component
 - ◆ $\mathcal{R}_0 = pcD = \sigma\tau cD$,
 - ◆ where σ measures intrinsic susceptibility and τ intrinsic tendency to transmit.
 - ◆ Why do we assume this is multiplicative?

Heterogeneity in TB

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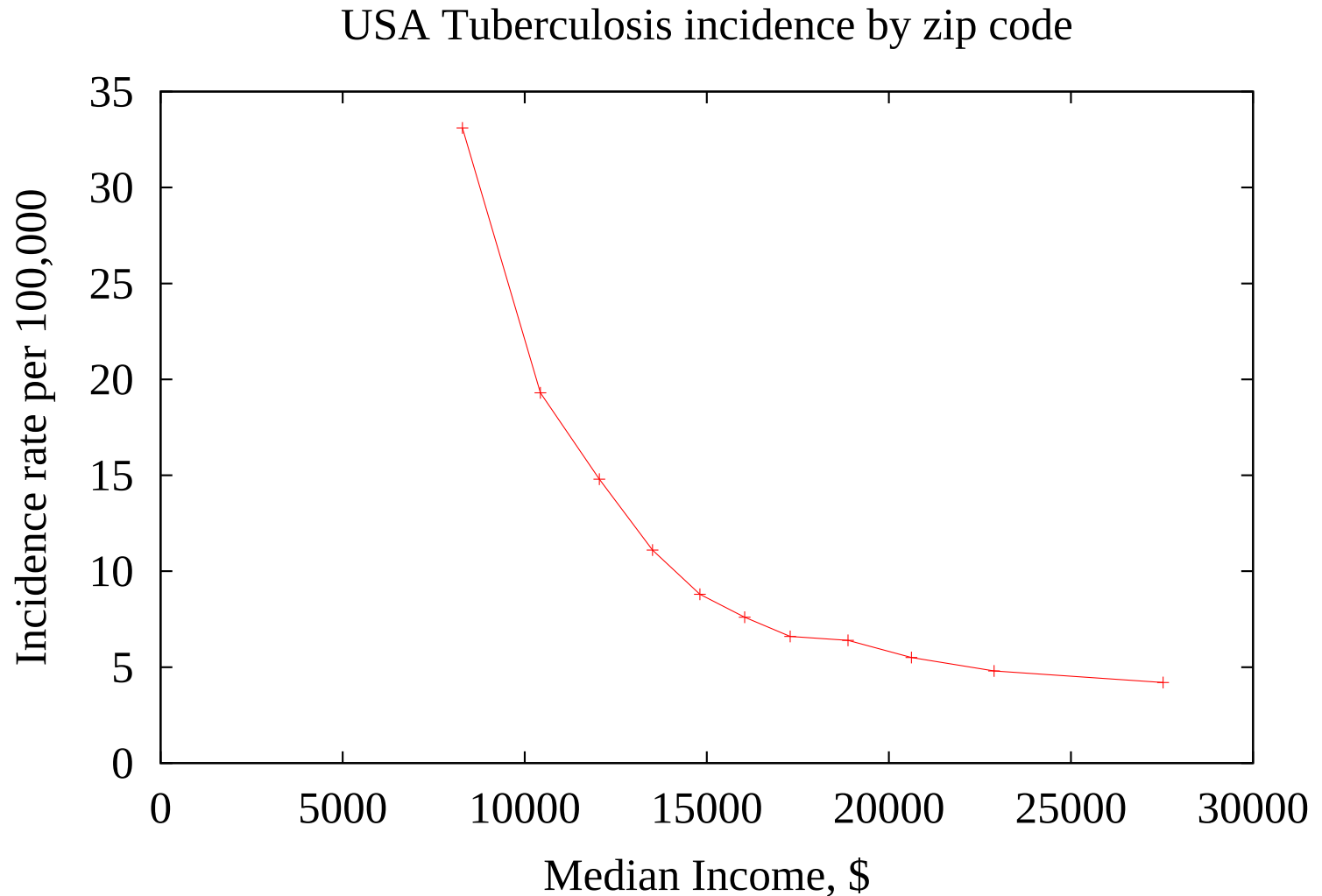
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Disease Heterogeneity in TB

■ *Progression to Disease*: Nutrition, stress

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- *Progression to Disease*: Nutrition, stress
- *Contact Rate*: Overcrowding, poor ventilation

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- *Progression to Disease*: Nutrition, stress
- *Contact Rate*: Overcrowding, poor ventilation
- *Cure Rate*: Access to medical care

Heterogeneity in other diseases

- *STDs*: Sexual mixing patterns, access to medical care

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Heterogeneity in other diseases

- *STDs*: Sexual mixing patterns, access to medical care
- *Influenza*: Crowding, nutrition

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- *STDs*: Sexual mixing patterns, access to medical care
- *Influenza*: Crowding, nutrition
- *Malaria*: Attractiveness to biting insects, geographical location, immune status

Heterogeneity in other diseases

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- *STDs*: Sexual mixing patterns, access to medical care
- *Influenza*: Crowding, nutrition
- *Malaria*: Attractiveness to biting insects, geographical location, immune status
- *Many others*

Parametric heterogeneity

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Imagine an aspect space indexed by $a \in \mathcal{A}$. Each a is associated with a value for $\sigma(a)$, $\tau(a)$, $c(a)$ and $D(a)$. We can model heterogeneity explicitly, using boxes, or distributions, or seek approximations based on the moments of the parameters: e.g., means, variances and covariances across the aspect space.

$$\bar{c} = \int_{\mathcal{A}} c(a) N(a) da$$

$$\sigma_c^2 = \int_{\mathcal{A}} (c(a) - (\bar{c}))^2 N(a) da$$

Mixing function

- $p(a, b)$ gives the proportion of group a 's contacts that are with members of group b .

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- $p(a, b)$ gives the proportion of group a 's contacts that are with members of group b .

- $$\int p(a, b)db = 1$$

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■ $p(a, b)$ gives the proportion of group a 's contacts that are with members of group b .

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$$c(a)N(a)p(a, b) = c(b)N(b)p(b, a)$$

Mixing function

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■ Simplest case is proportional mixing:

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- Simplest case is proportional mixing:

- ◆
$$p(a, b) = \frac{c(b)N(b)}{\bar{c}N}$$

Next-generation framework

- We can analyze the relationship between risk factors and equilibria without making assumptions about waiting times, using a next-generation framework:

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- We can analyze the relationship between risk factors and equilibria without making assumptions about waiting times, using a next-generation framework:
- *How many new cases are caused by a given case?*

Next-generation framework

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- We can analyze the relationship between risk factors and equilibria without making assumptions about waiting times, using a next-generation framework:
- *How many new cases are caused by a given case?*
 - ◆ $\mathcal{R}_0$ is the expected number of new cases per case along the eigendistribution of cases near the disease-free equilibrium.

Next-generation framework

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- *How many new cases are caused by a given case?*
 - ◆ $\mathcal{R}_0$ is the expected number of new cases per case along the eigendistribution of cases near the disease-free equilibrium.
 - ◆ The endemic equilibrium is where a distribution of cases regenerates itself for the next generation.

Heterogeneity and $\mathcal{R}_0$

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Proportional mixing

$$\mathcal{R}_0 = \frac{\langle c^2 \sigma \tau D \rangle_N}{\langle c \rangle_N}$$

- Heterogeneity in susceptibility, transmissibility or duration alone has no effect on $\mathcal{R}_0$ and thus on whether a disease persists in a population.

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- Heterogeneity in contact rate alone increases $\mathcal{R}_0$. The “effective” contact rate is given by $c_e = \bar{c}(1 + V_c^2)$, where V_c is the coefficient of variation of the distribution of c .

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- Correlations between other parameters can have a similar effect (e.g., if those who are more susceptible are on average also more able to transmit).

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Assortative mixing

- Assortative mixing means the tendency of like to mix with like

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Assortative mixing

- Assortative mixing means the tendency of like to mix with like
- Assortative mixing combined with any kind of heterogeneity will increase $\mathcal{R}_0$.

Heterogeneity and equilibrium

- If a disease persists, in the long term the geometric mean reproductive number $\langle \mathcal{R} \rangle = 1$

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- If a disease persists, in the long term the geometric mean reproductive number $\langle \mathcal{R} \rangle = 1$
- The factors that reduce $\mathcal{R}$ from $\mathcal{R}_0$ to 1 determine the level of equilibrium.

Heterogeneity and equilibrium

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 - ◆ If the only factor that changes is S/N , we have the classic endemic curve

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- In a heterogeneous population, the infected population and the susceptible population generally both become less risky on average as the disease spreads!

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 - ◆ Makes equilibrium disease level lower for a given risk

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- In a heterogeneous population, the infected population and the susceptible population generally both become less risky on average as the disease spreads!
 - ◆ Makes equilibrium disease level lower for a given risk
 - ◆ Conversely, risk may be higher than you think

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Proportional mixing

$$\blacksquare \mathcal{R}_0 = \langle c\sigma \rangle_N \frac{\langle c\tau D \rangle_{\hat{I}}}{\bar{c}}; \quad 1 = \langle c\sigma \rangle_{\tilde{S}} \frac{\langle c\tau D \rangle_{\tilde{I}}}{\bar{c}} \frac{S}{N}$$

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- Variation in transmissibility alone has no effect on average transmission, or on the endemic curve.
- Variation in susceptibility alone reduces transmission at equilibrium, thus reduces level of disease (flattens the curve).
- Variation in contact rate alone reduces transmission at equilibrium compared to during invasion (flattens the curve).

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- Variation in susceptibility alone reduces transmission at equilibrium, thus reduces level of disease (flattens the curve).
- Variation in contact rate alone reduces transmission at equilibrium compared to during invasion (flattens the curve).
 - ◆ Effect on equilibrium level of disease is mixed.

Homogeneous population

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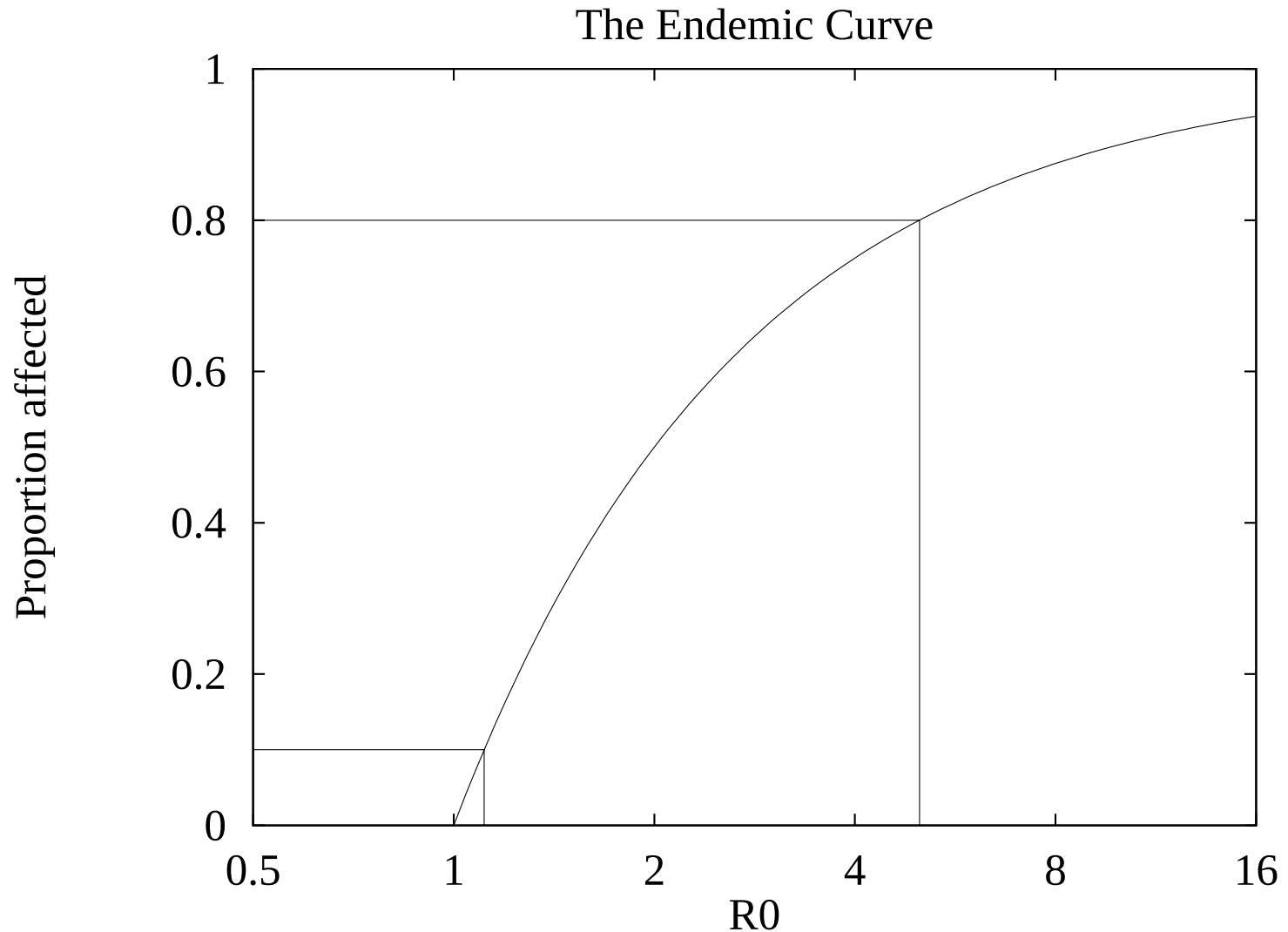
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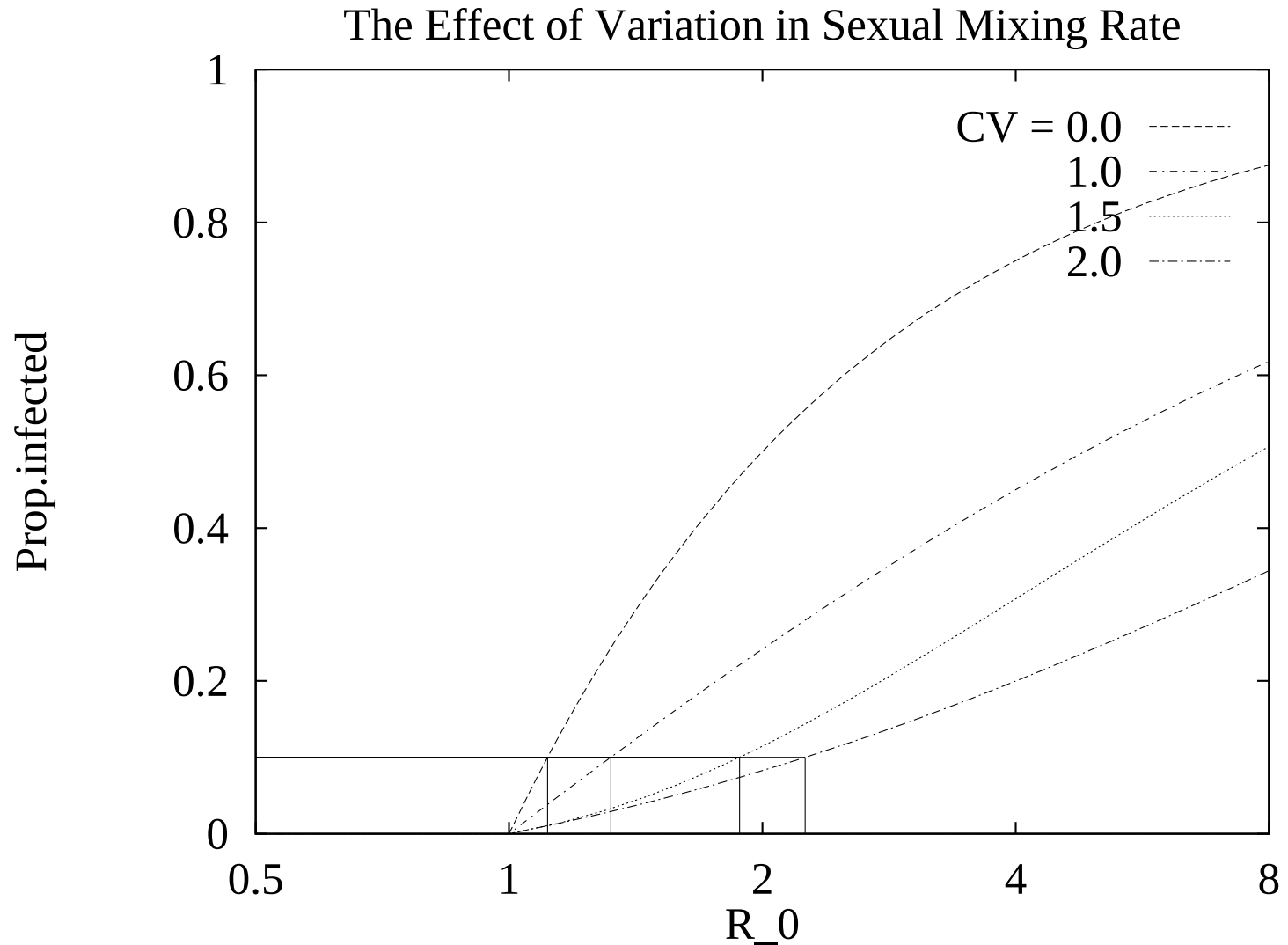
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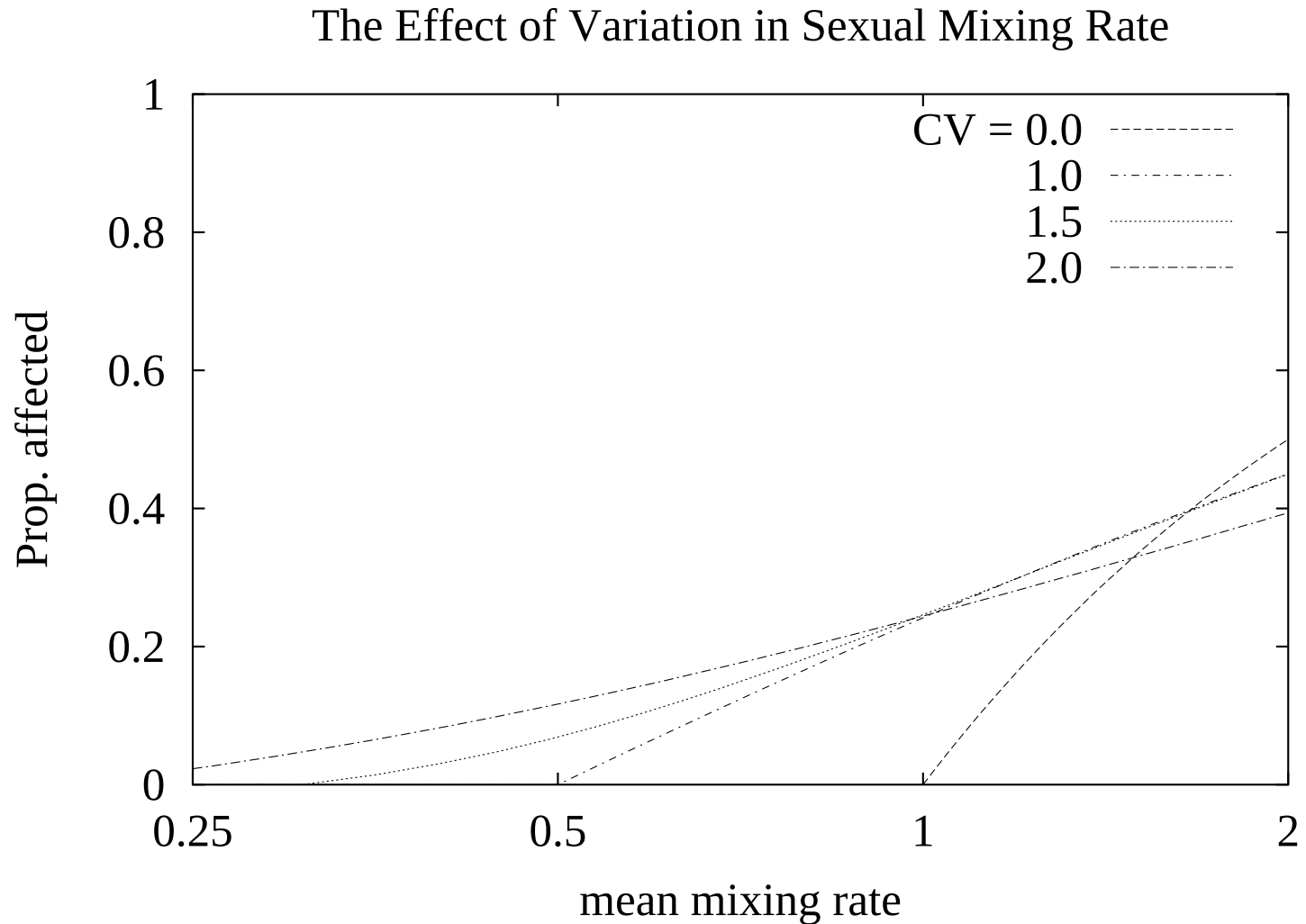
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Approximations based on moments

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The effects of heterogeneity

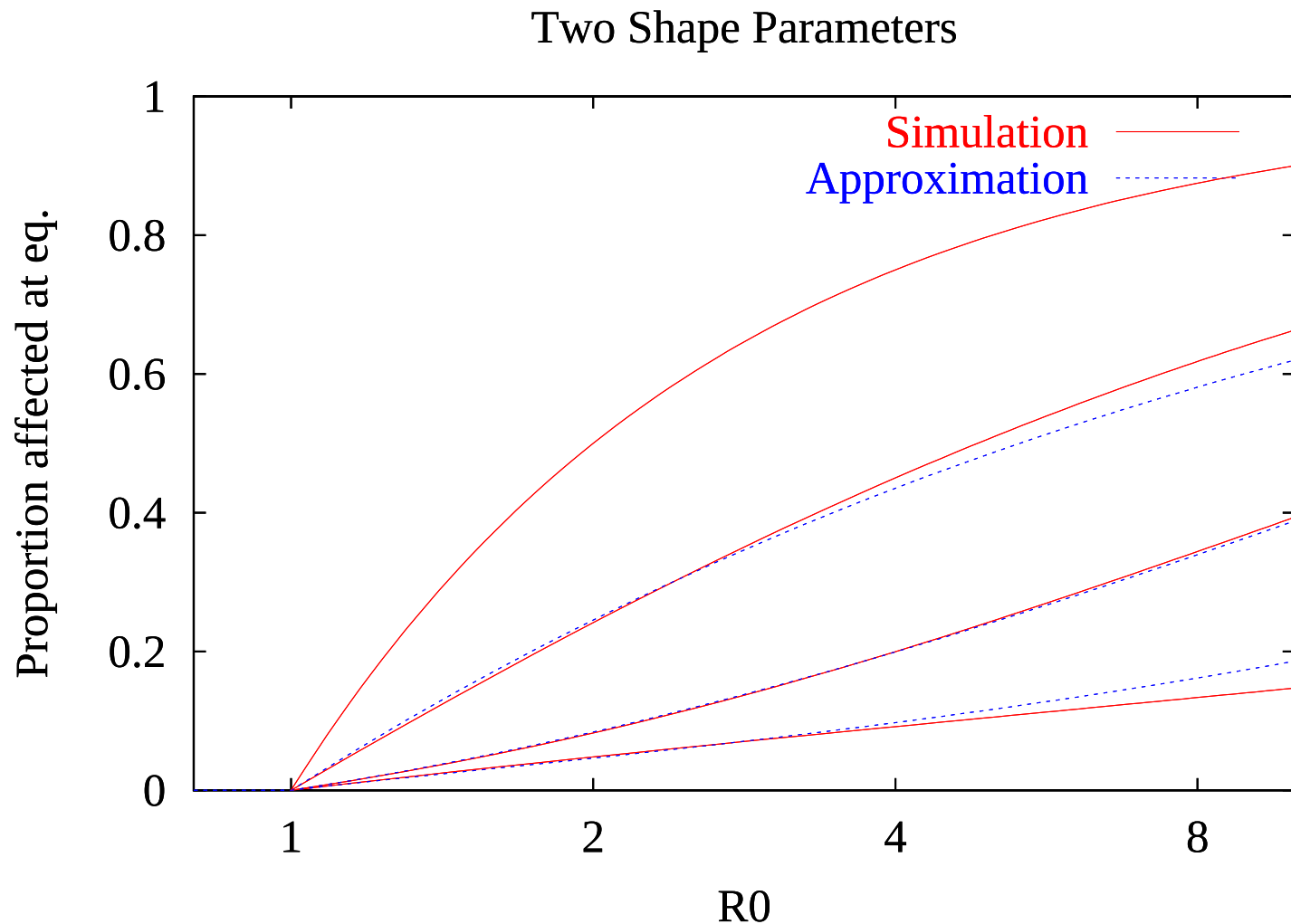
● Disease Heterogeneity in TB

● Heterogeneity in other diseases

● Next-generation framework

● Heterogeneity and $\mathcal{R}_0$

● Heterogeneity and equilibrium



Host heterogeneity summary

- Heterogeneity makes diseases harder to eliminate than would be predicted from the basic model

- Population heterogeneity

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 - ◆ Assortative mixing
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- Disease-control strategies should take heterogeneity on a variety of scales into account.
- Simple moment-based approximations can help extend the usefulness of the endemic curve – relationships between risk factors and disease level – to heterogeneous populations.