

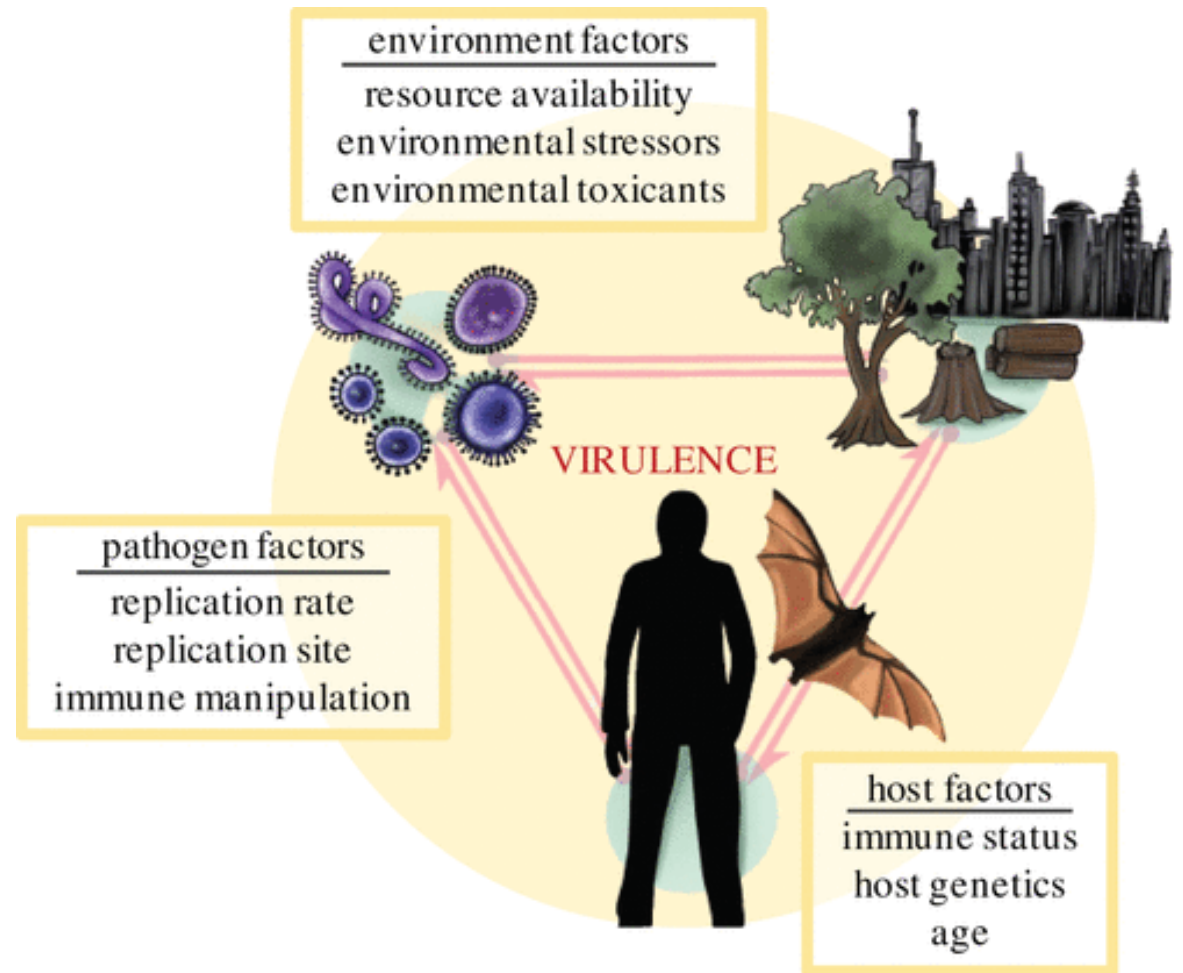
The three Ts of virulence evolution during zoonotic emergence

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March 15, 2022

What do we mean by virulence?

In disease ecology, virulence is defined as the additional mortality rate due to infection in mathematical models.

For virulence evolution, we are primarily interested in the pathogen factors controlling disease severity, though host and environmental factors will also affect infection outcomes.



Visher et al. (2021). PRSB.

**Can we
even predict
pathogen
evolution?**



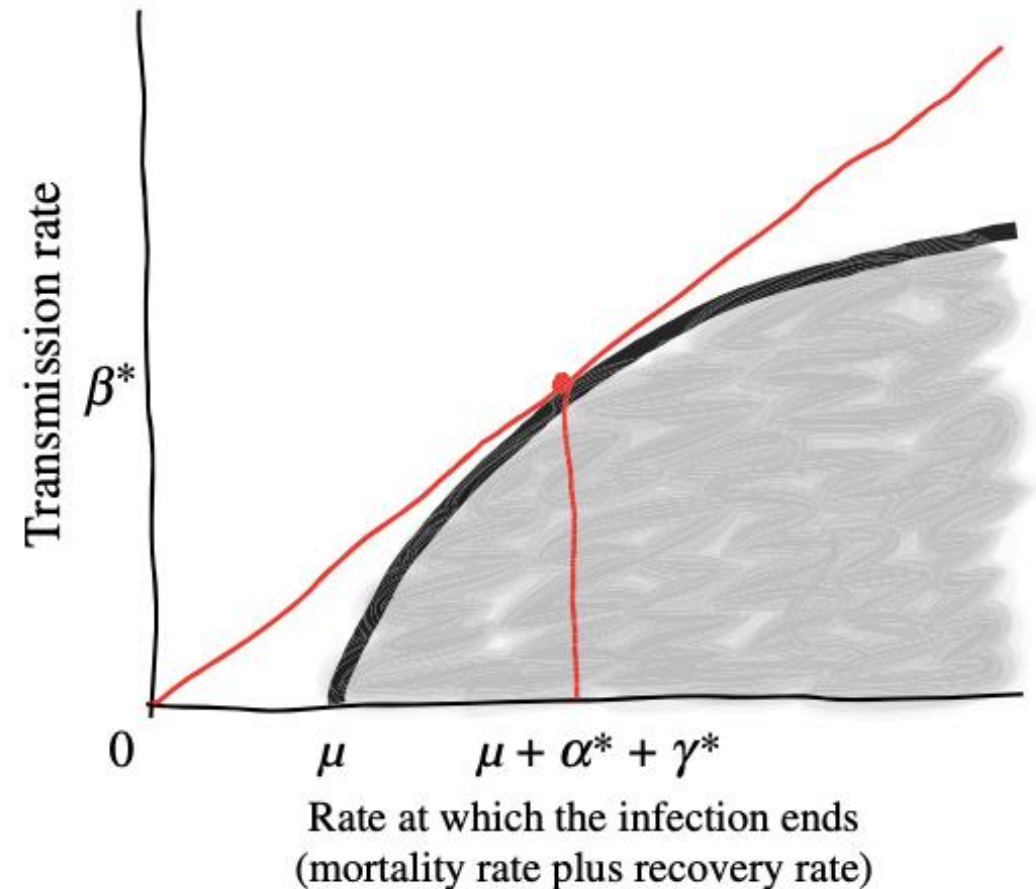
Virulence can adaptively evolve

*Myxoma virus
in rabbits*



The Virulence – Transmission Trade-off

Modern theory generally predicts that pathogens “should” evolve **intermediate** levels of virulence because the costs of virulence trade-off with the benefits of higher transmission rates.

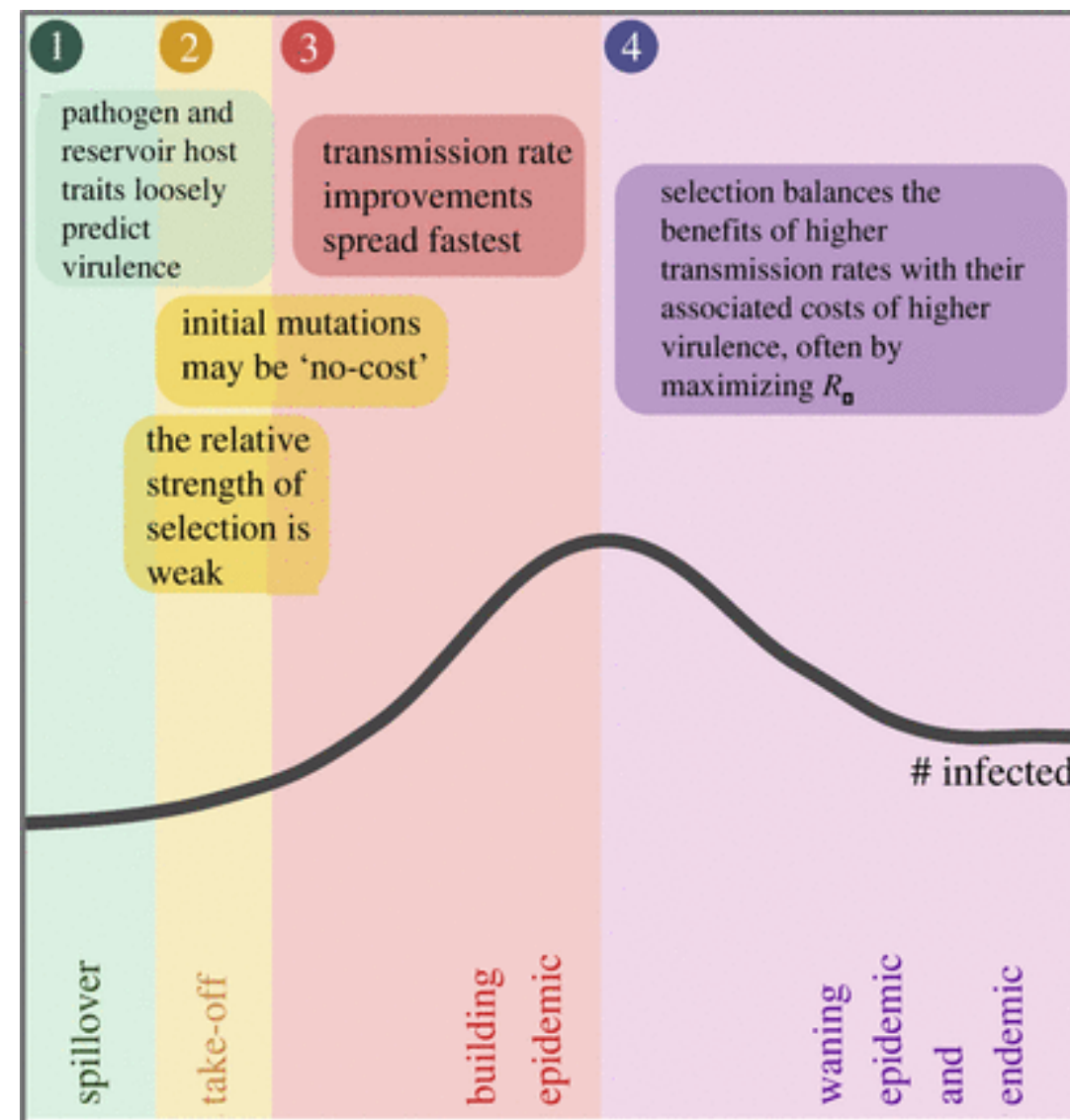


**How does
this change
in newly
emerged
zoonotic
diseases?**

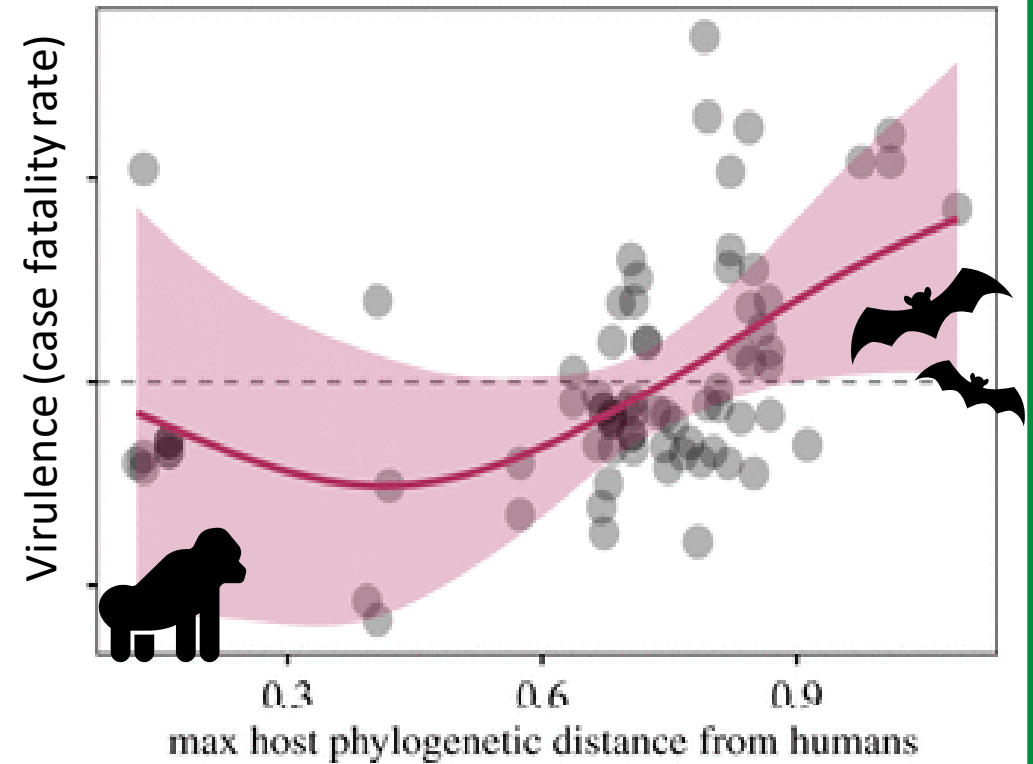
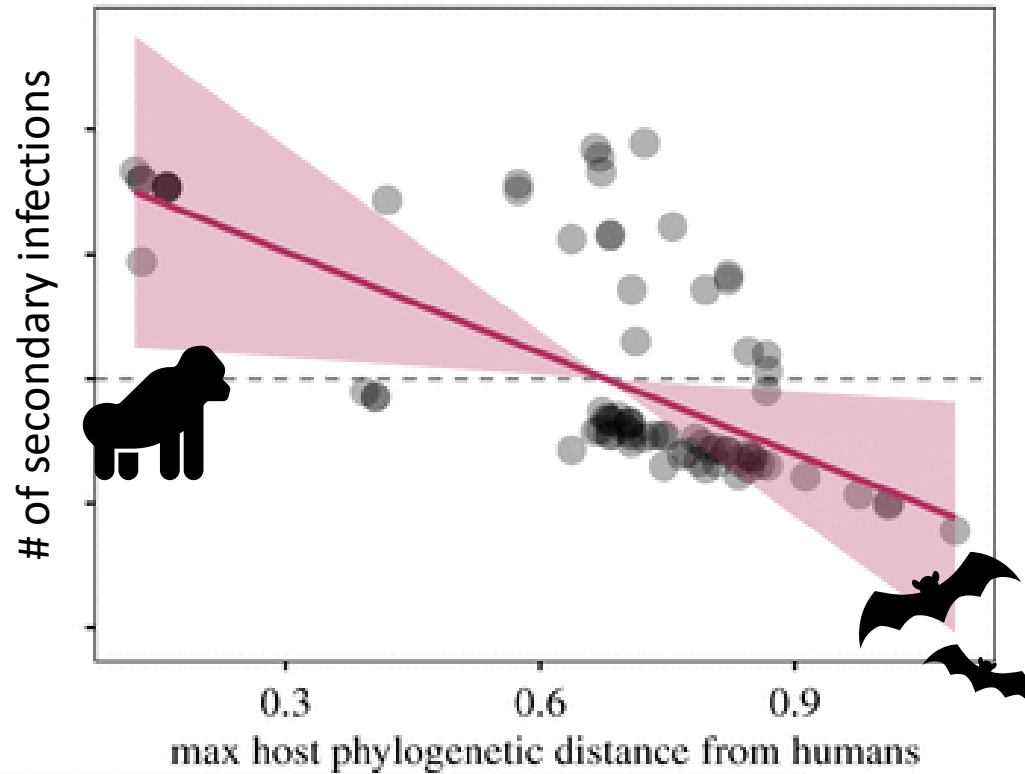


The three Ts:

1. Trade-Offs
2. Transmission
3. Time Scales



Pathogen and reservoir host traits loosely predict R_0 and virulence



Stochastic effects in small populations can overwhelm selection

1

pathogen and reservoir host traits loosely predict virulence

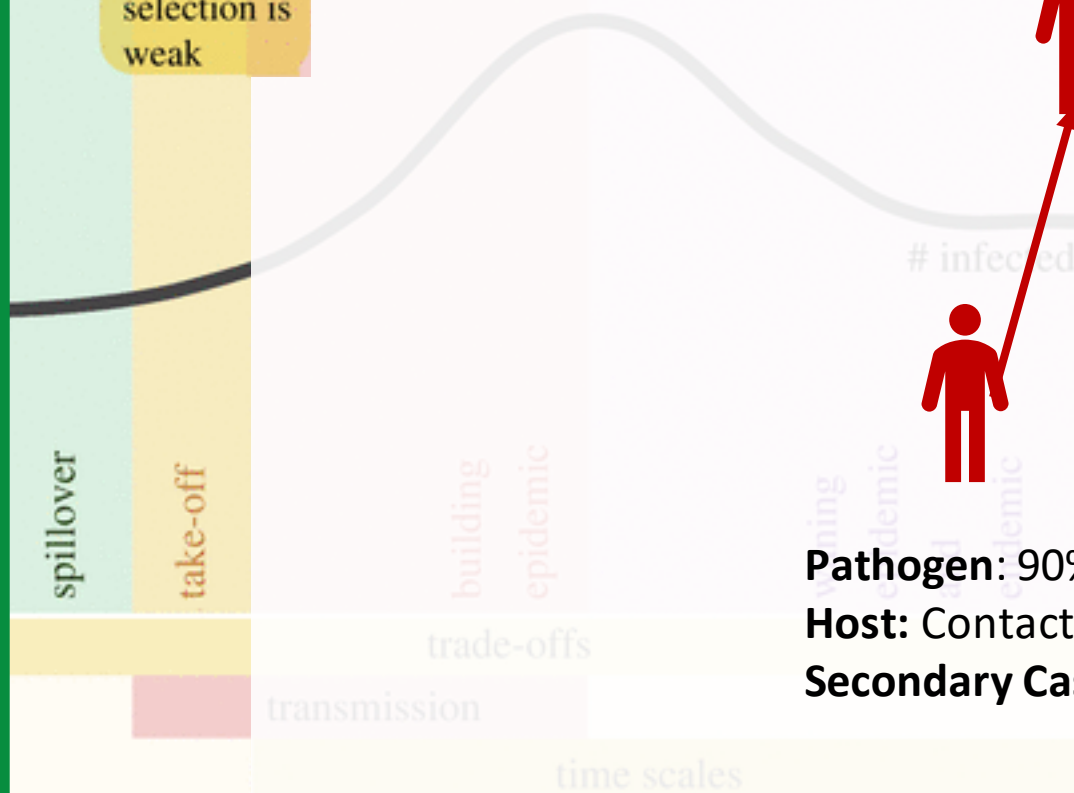
2

initial mutations may be 'no-cost'

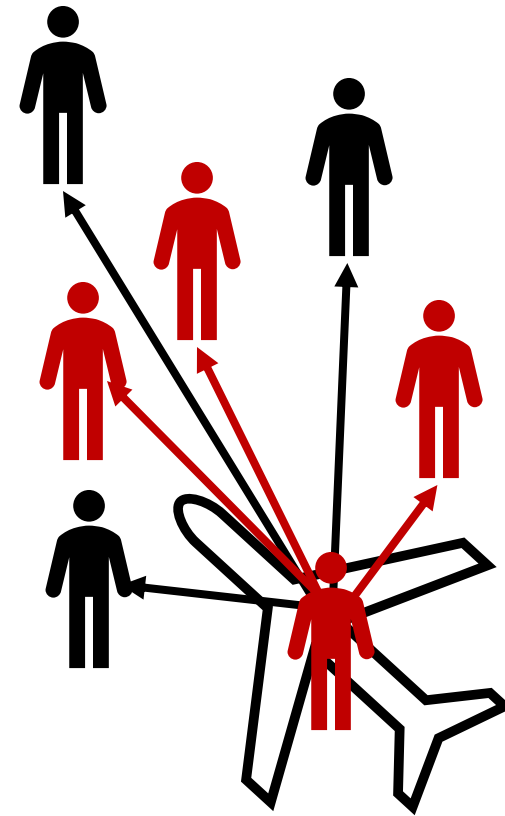
the relative strength of selection is weak

4

selection balances the transmission rates with the associated costs of higher virulence, often by maximizing R_0

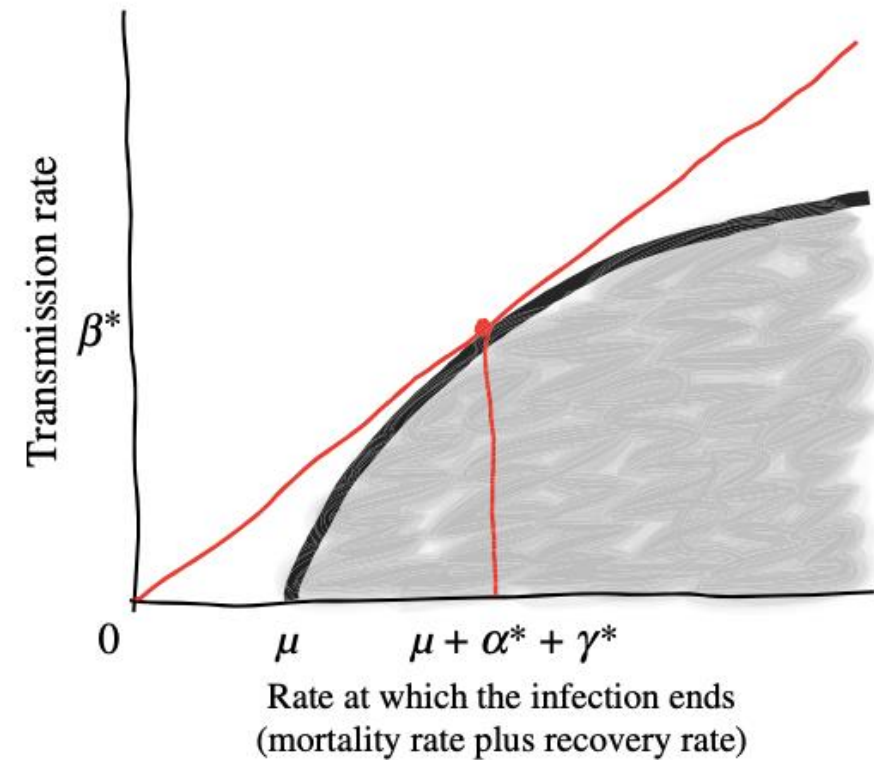


Pathogen: 90% infection rate
Host: Contacts 1 Susceptible
Secondary Cases = 1

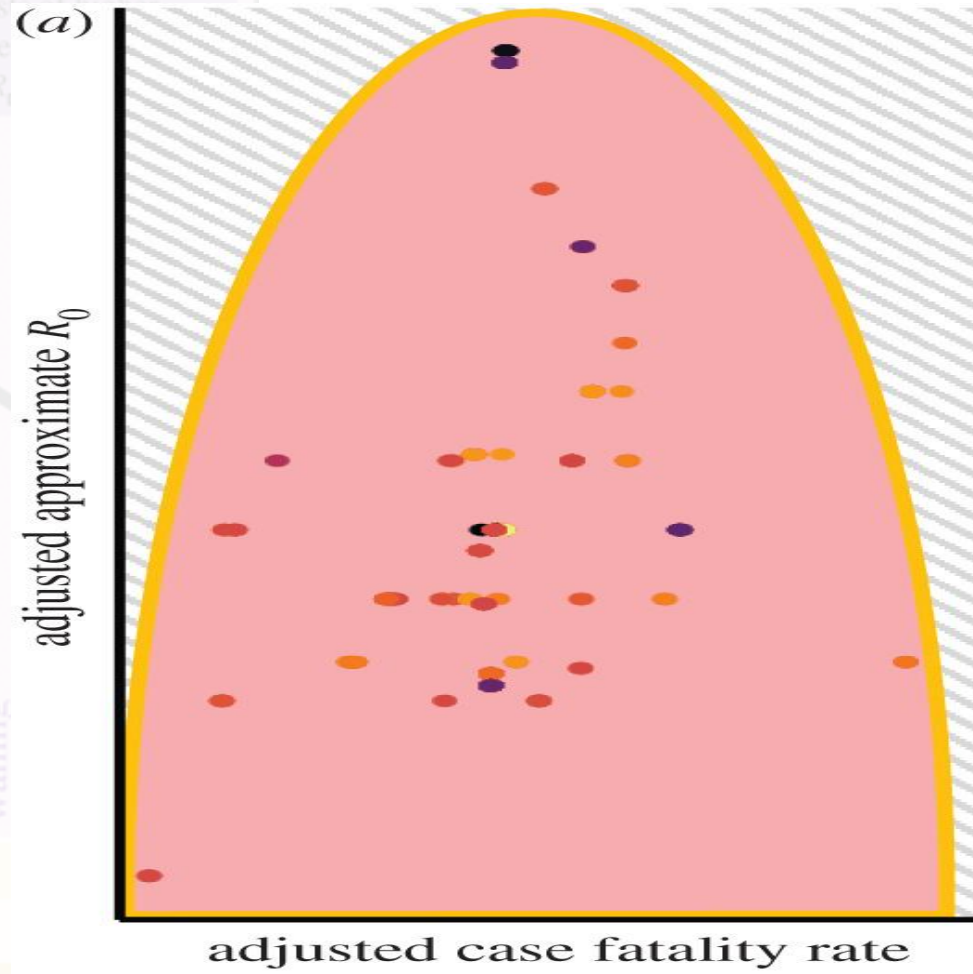
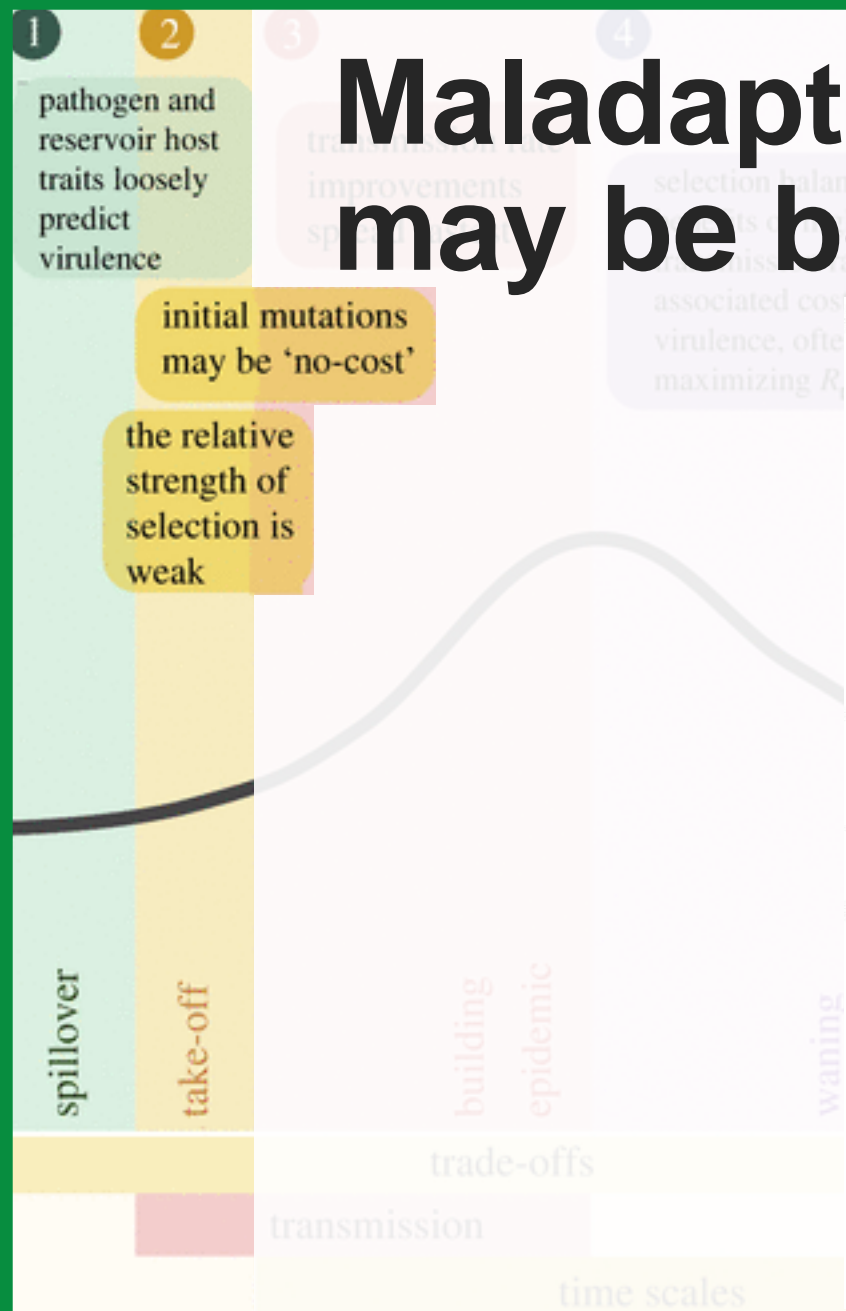


Pathogen: 50% infection rate
Host: Contacts 6 Susceptibles
Secondary Cases = 3

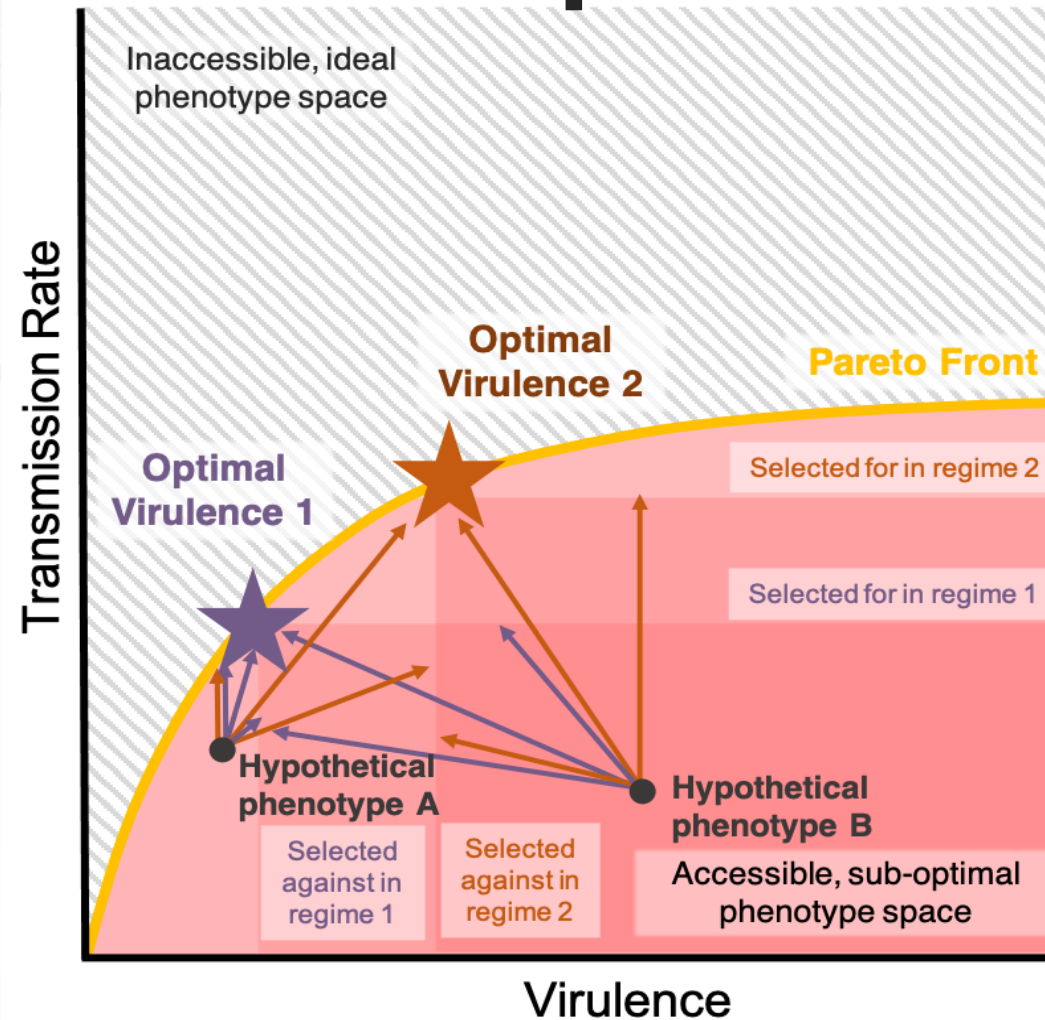
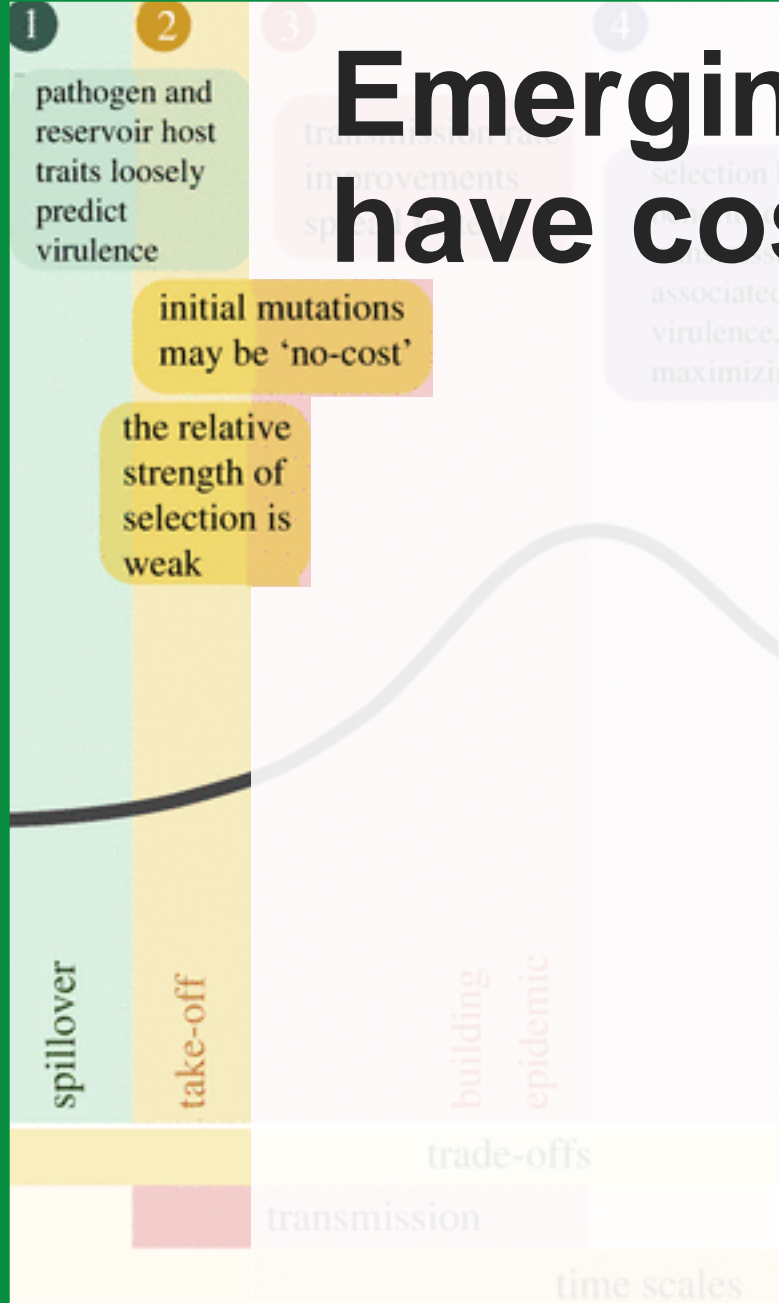
Why is it difficult to predict how a novel zoonotic pathogen will evolve when it spills over into humans?

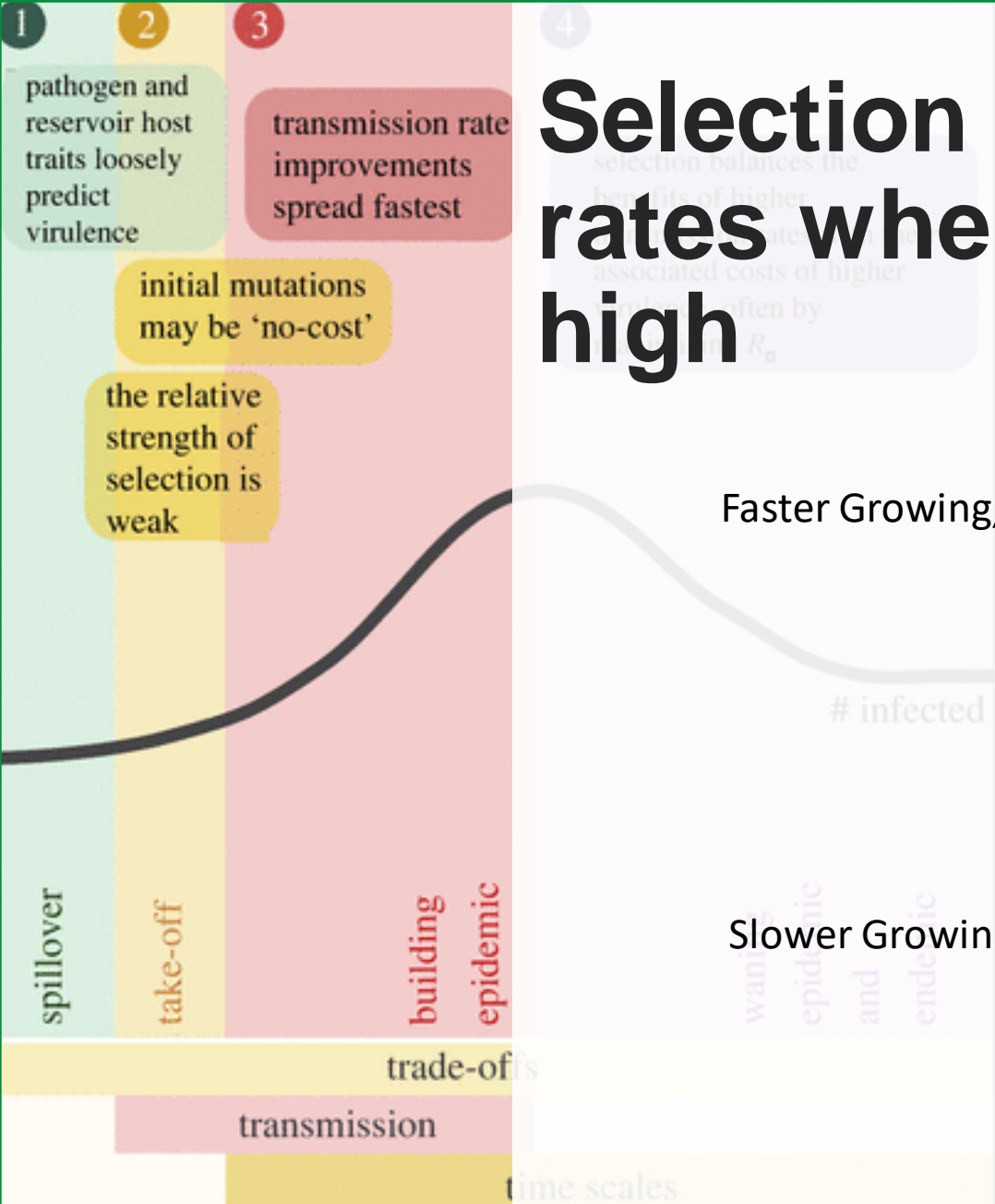


Maladapted zoonotic pathogens may be below the Pareto front



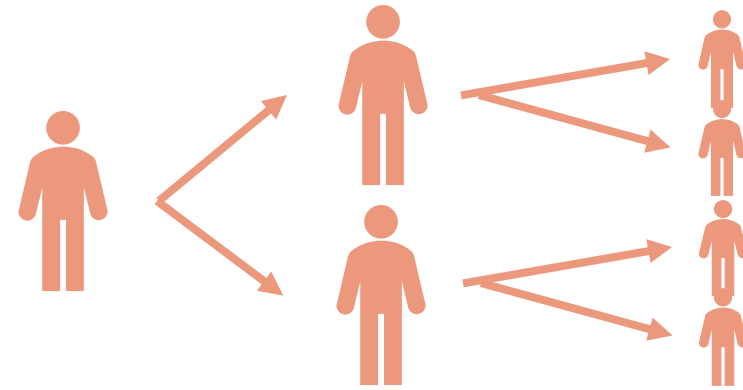
Emerging zoonotic pathogens may have costless adaptive mutations



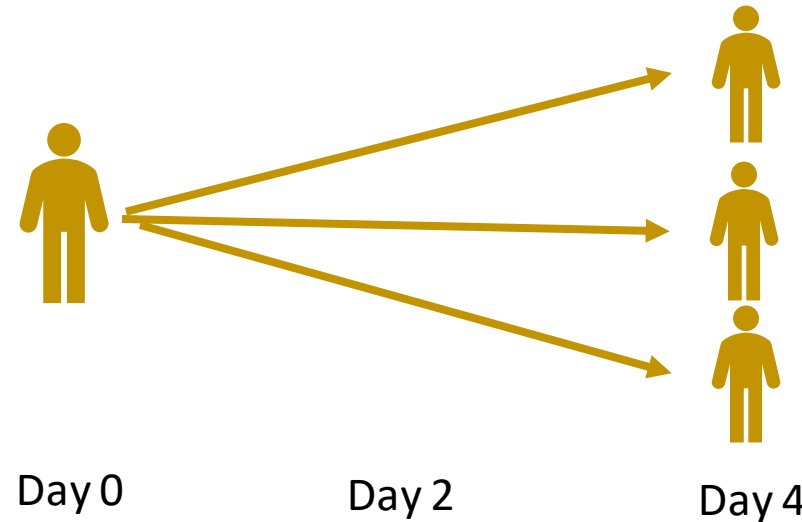


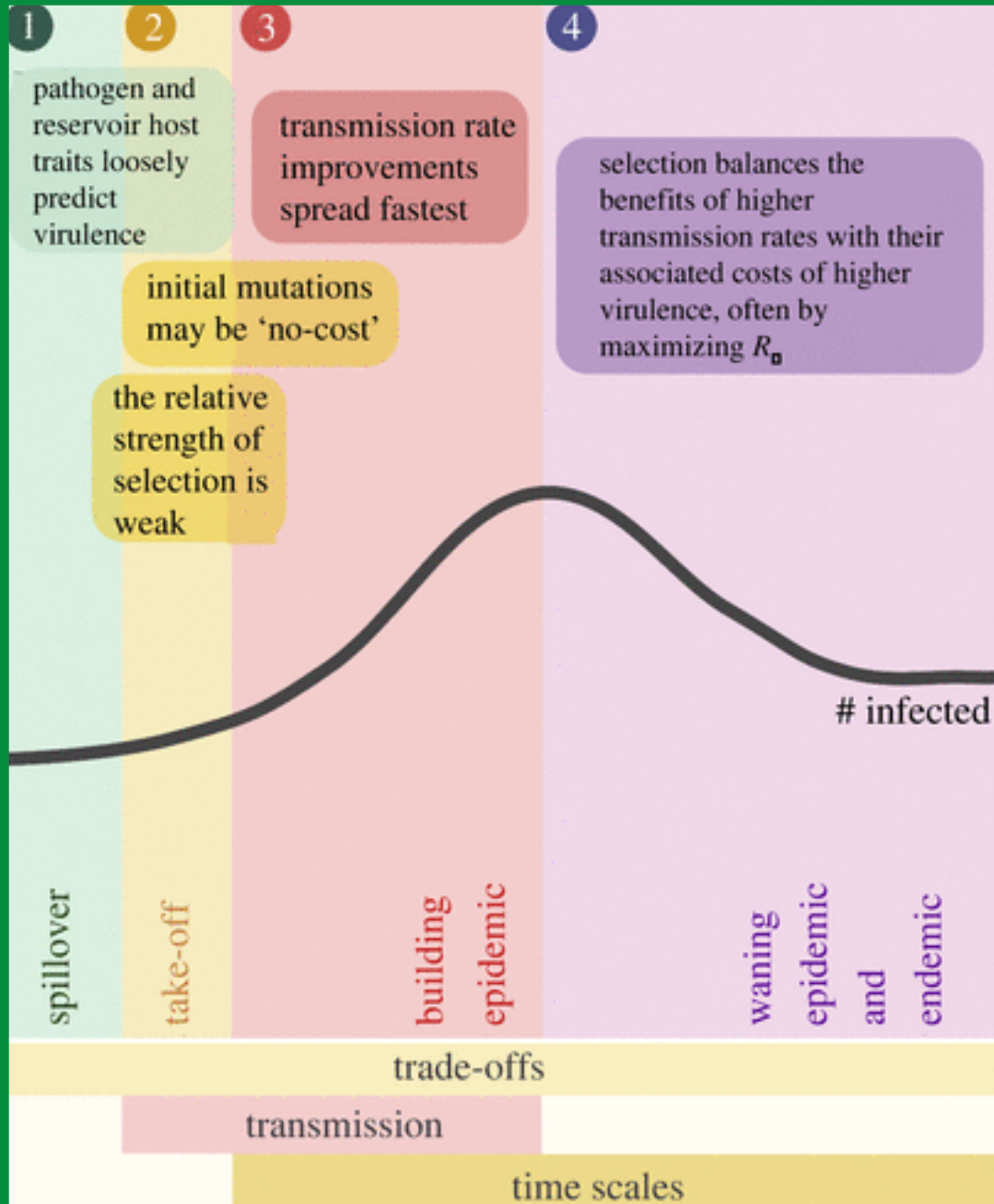
Selection favors high transmission rates when susceptible density is high

Faster Growing, Lower R_0

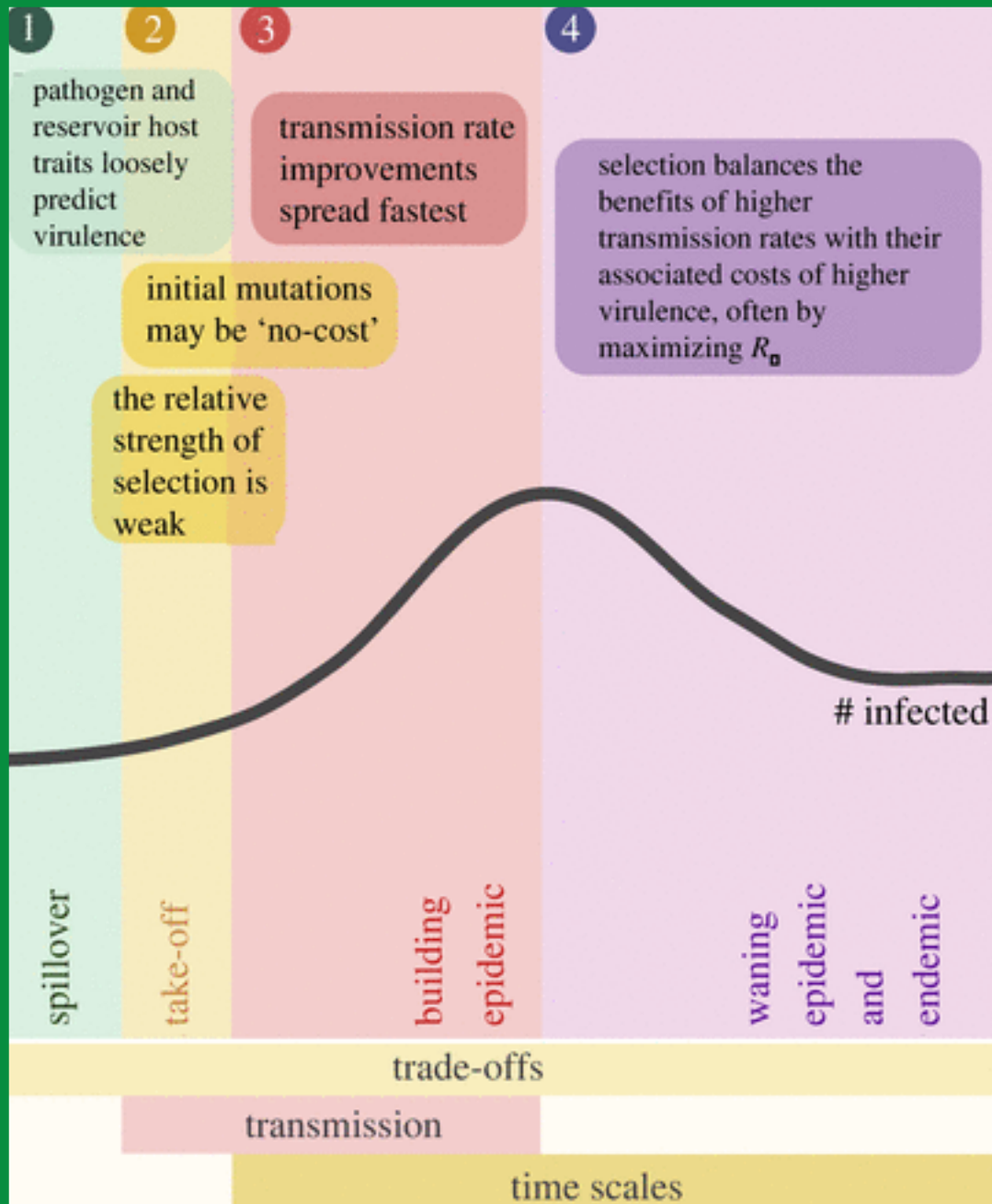


Slower Growing, Higher R_0





Selection favors high R_0
 (the total number of secondary infections in a susceptible population) **at equilibrium**



Conclusions...

Virulence and transmission rate may trade-off

Selection may increase transmission rate

Evolutionary dynamics may change with time

Can we evolutionarily manage virulence?



PROCEEDINGS B

royalsocietypublishing.org/journal/rspb

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