

Modelling Fitness Trade-Offs of Rates of Horizontal Gene Transfer

Evolution 02-731 Project

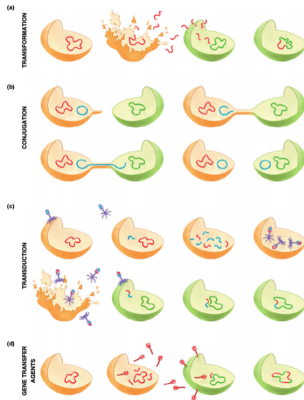
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Background

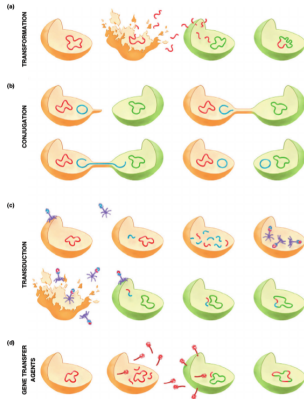
Horizontal Gene Transfer (HGT)



- **Transformation:**
Incorporation of free-floating DNA into the genome

Figure 1: HGT Mechanisms

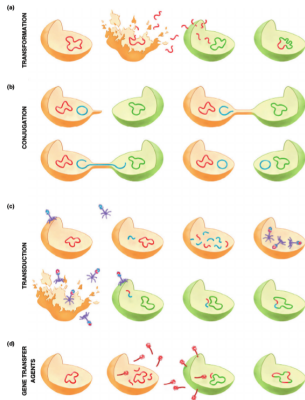
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- **Transduction:** Transfer of DNA via phage

Figure 1: HGT Mechanisms

CRISPR-Cas In Bacteria

► Adaptive Bacterial Immune System

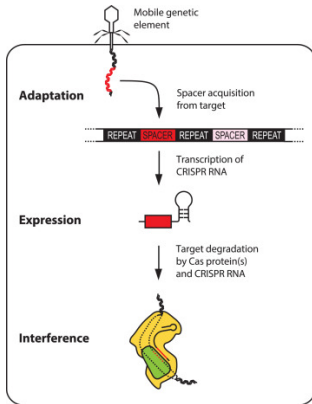
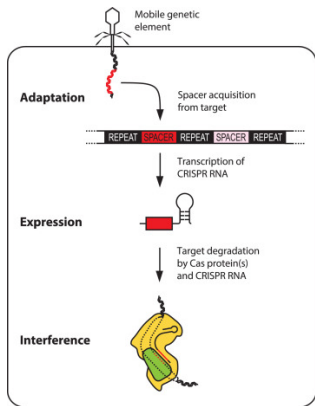


Figure 2: CRISPR-Cas Mechanism

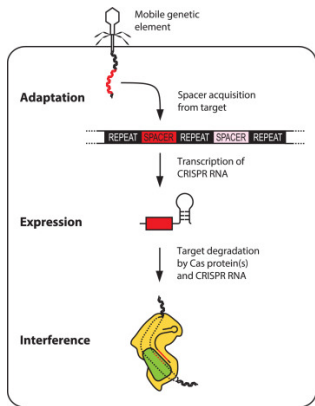
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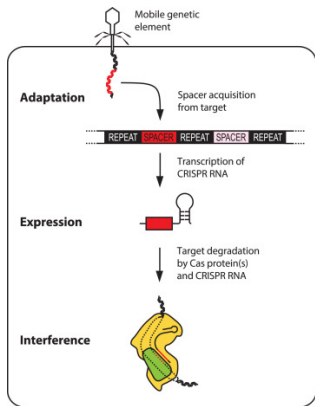
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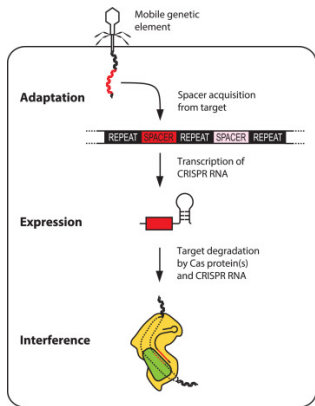
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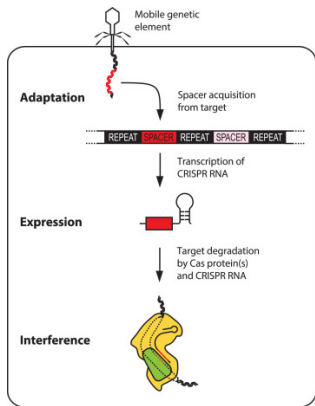
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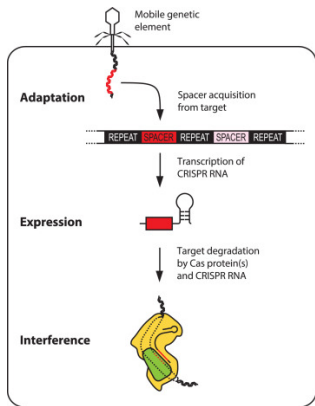
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- ▶ Adaptive Bacterial Immune System
- ▶ Requires CRISPR array and Cas proteins
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 1. Exposure
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 3. Targeted degradation in next exposure
- ▶ Protects against “foreign” DNA, but can acquire **any** DNA as a spacer

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- ▶ HGT can result in genome disruption or gaining toxic gene products
- ▶ metabolic cost to maintain CRISPR or HGT machinery expression
- ▶ CRISPR systems can also be transferred between bacteria via plasmids

Model

Genotypes

Allele		Description
Major	Minor	
<i>R</i>	<i>r</i>	has/does not have resistance gene
<i>H</i>	<i>h</i>	HGT machinery is expressed/not expressed
<i>C</i>	<i>c</i>	CRISPR-Cas is expressed/not expressed

Table 1: Allele definitions

Basics

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- ▶ **Alternating Threat:** Same as Cyclical Threat model but switching between threats each event.

Fitness

<i>Genotype</i>	<i>Environment</i>		
	E_n	E_b	E_a
<i>RCH</i>	$1 - 2s_m$	$(1 + s_p)(1 - 2s_m)$	$(1 + s_p)(1 - 2s_m)$
<i>RCh</i>	$1 - s_m$	$(1 + s_p)(1 - s_m)$	$(1 + s_p)(1 - s_m)$
<i>RcH</i>	$1 - s_m$	$1 - s_m$	$(1 + s_p)(1 - s_m)$
<i>Rch</i>	1	1	$1 + s_p$
<i>rCH</i>	$1 - 2s_m$	$(1 + s_p)(1 - 2s_m)$	$1 - 2s_m$
<i>rCh</i>	$1 - s_m$	$(1 + s_p)(1 - s_m)$	$1 - s_m$
<i>rch</i>	$1 - s_m$	$1 - s_m$	$1 - s_m$
<i>rch</i>	1	1	1

Table 2: Relative fitness values for each genotype in each environment

► s_m reflects cost of maintaining HGT/CRISPR, $s_m \ll s_p$

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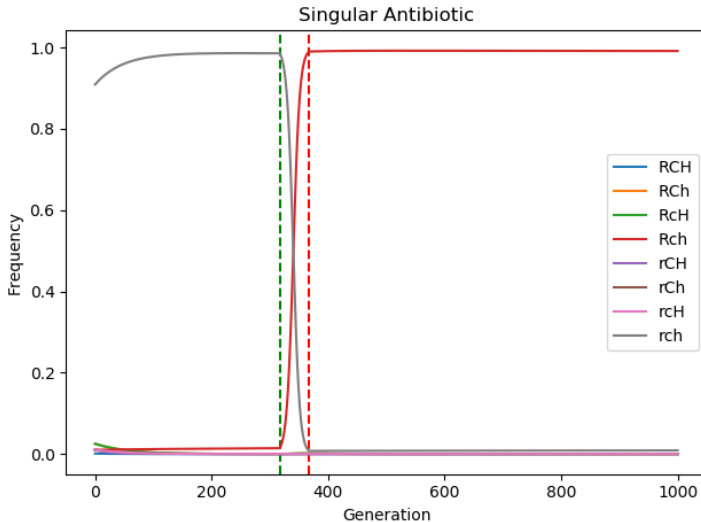
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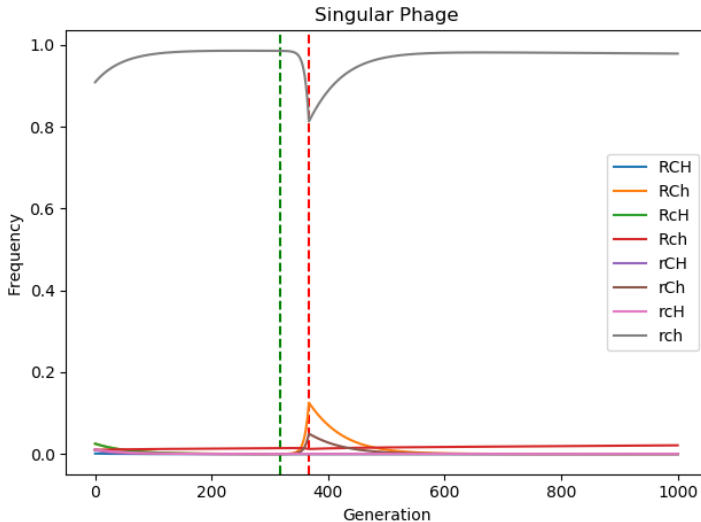
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Results

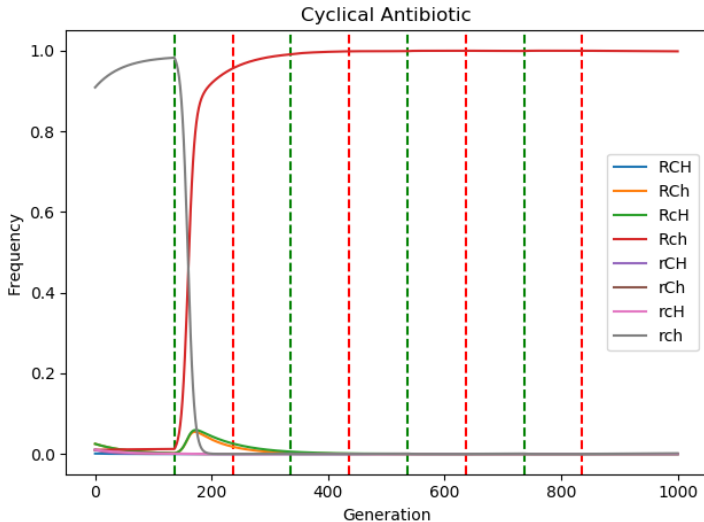
Single Antibiotic Event



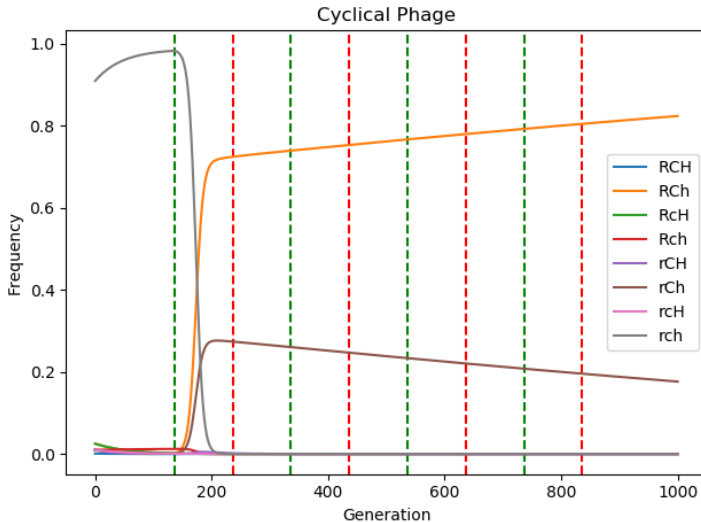
Single Phage Event



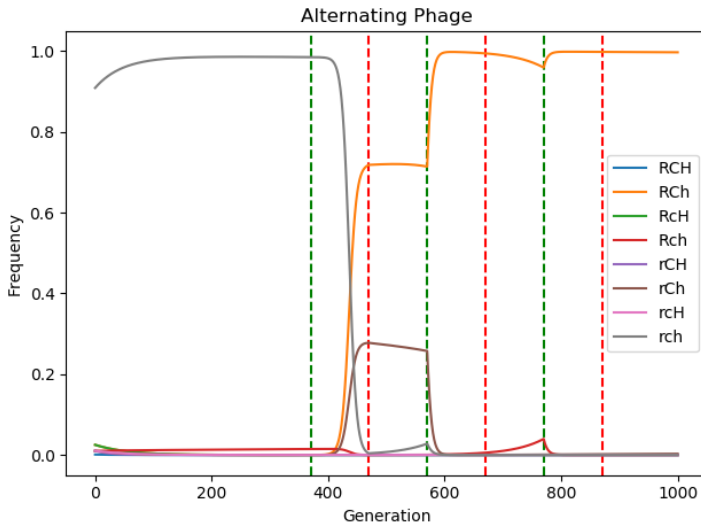
Cyclical Antibiotic Events



Cyclical Phage Events

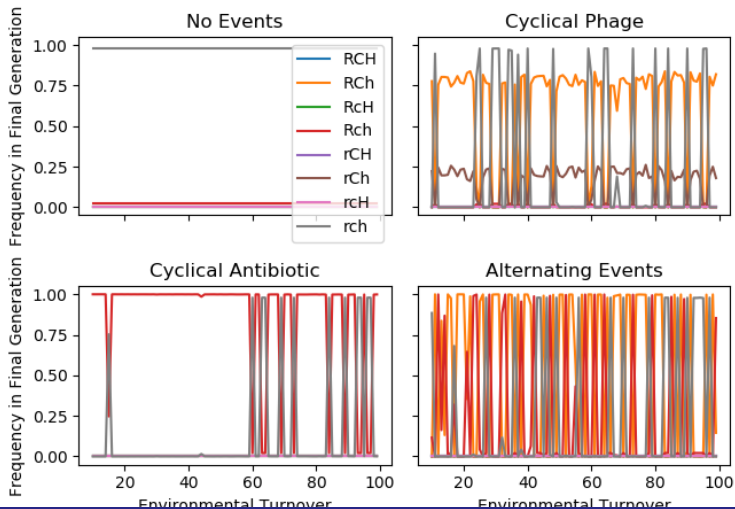


Alternating Events



Environmental Stability

Genotype Frequencies vs Environmental Turnover



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Future Directions

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- ▶ model phage population dynamics directly
- ▶ incorporate terms that reflect biological trade-off of HGT/CRISPR