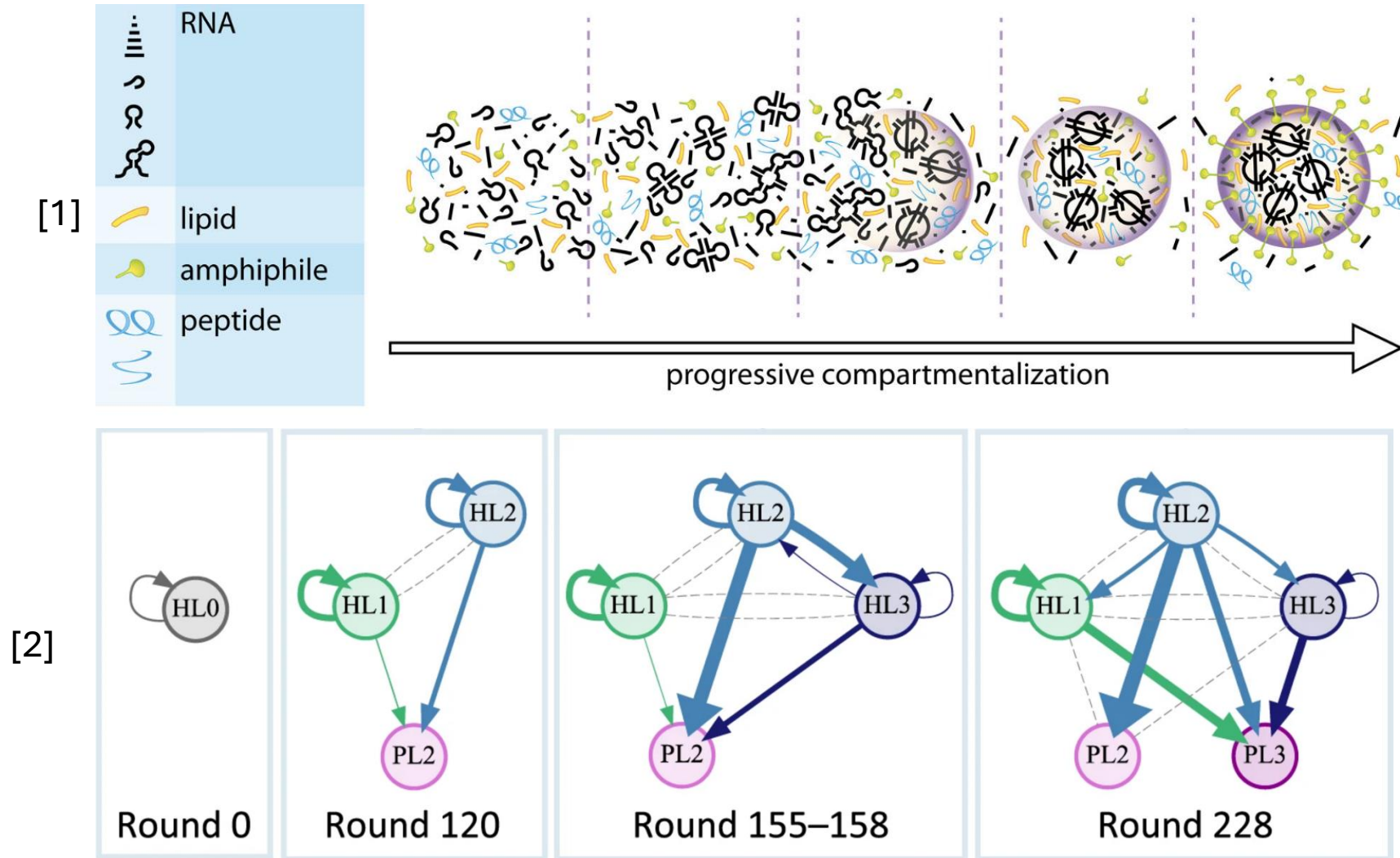


# Compositional memory matters for early molecular systems

Barnabe Ledoux

# Evolution of complexity



[1] P.Nghe, *A stepwise emergence of evolution in the RNA world*, FEBS Letters, 10.1002/1873-3468.70065 (2025)

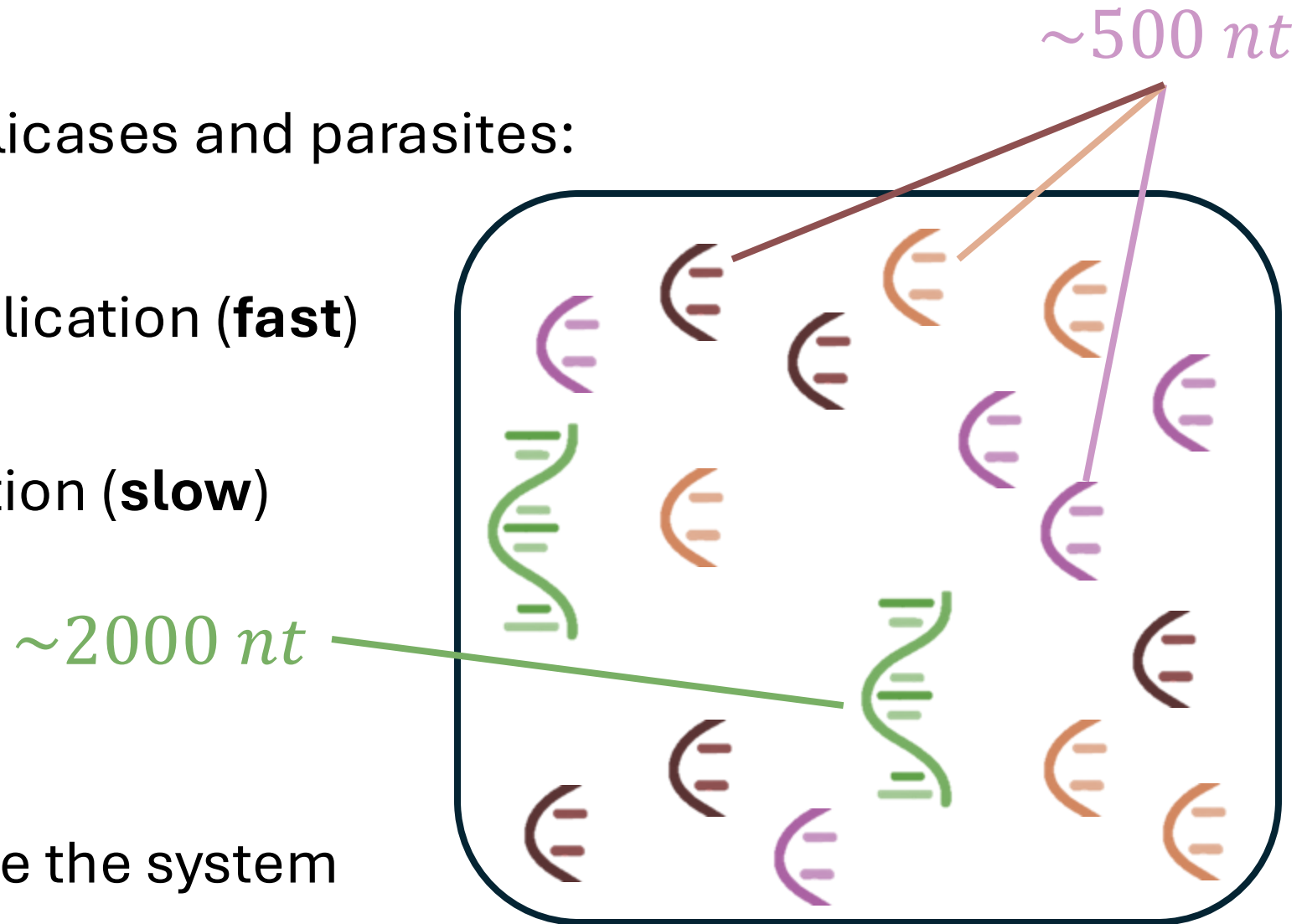
[2] R Mizuuchi, T Furubayashi, N Ichihashi, *Evolutionary transition from a single RNA replicator to a multiple replicator network*. Nat. Commun. 13, 1460 (2022)

# The need for compartments

In a **pool** with replicases and parasites:

- Parasite replication (**fast**)
- Self replication (**slow**)

⇒ Parasites invade the system

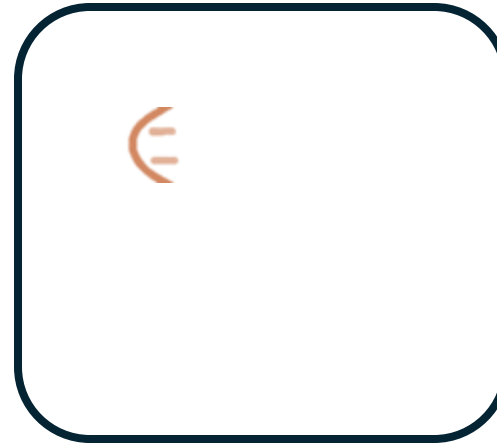


# The need for compartments

With compartments:

- Parasite replication (**fast**)
- Self replication (**slow**)
- Parasites alone cannot replicate

Isolated parasite

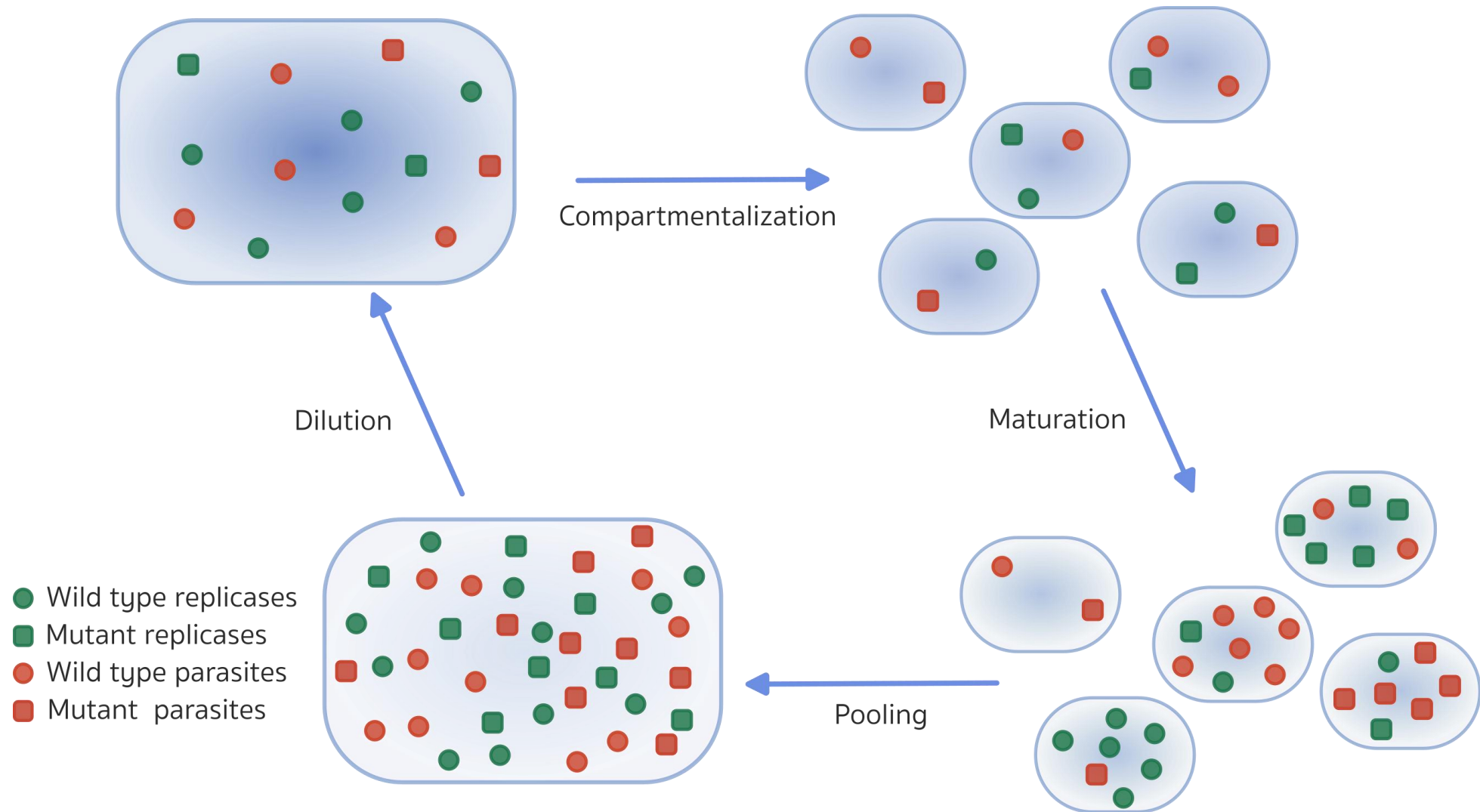


# Modeling dynamics

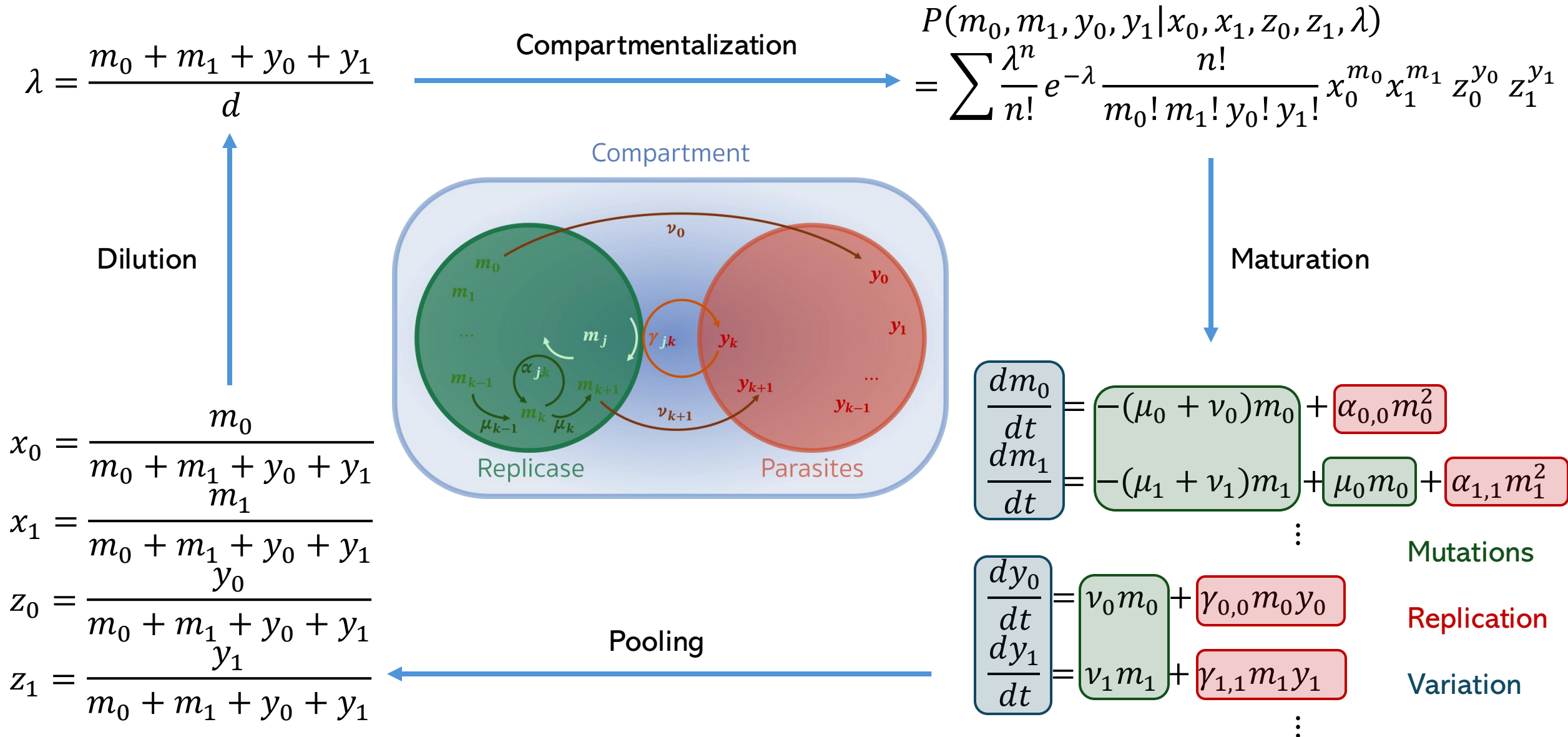
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# Transient compartmentalization with natural selection and mutations



# Evolution equations



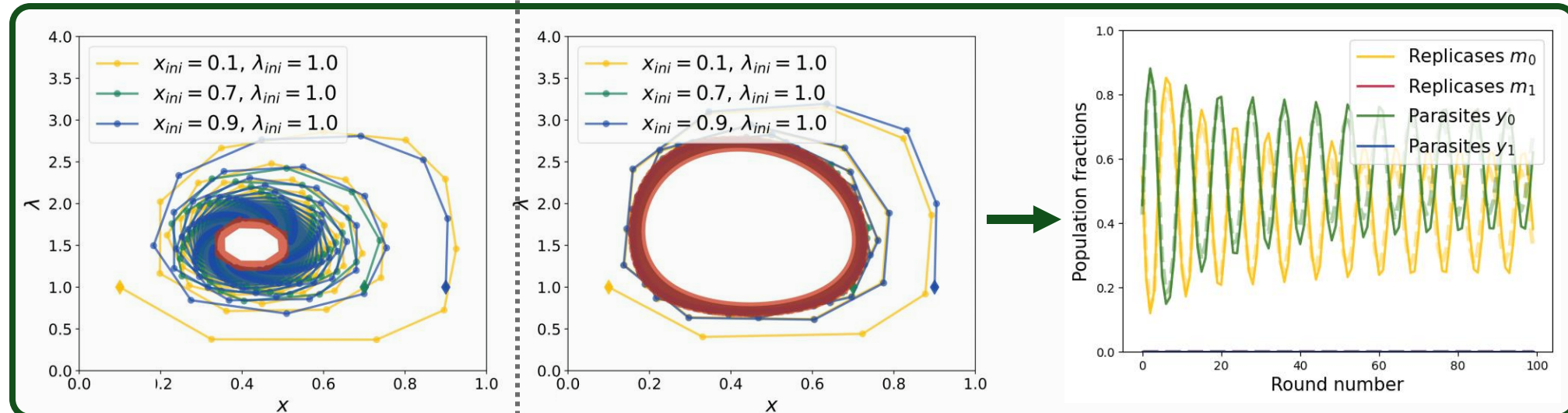


# Oscillatory dynamics

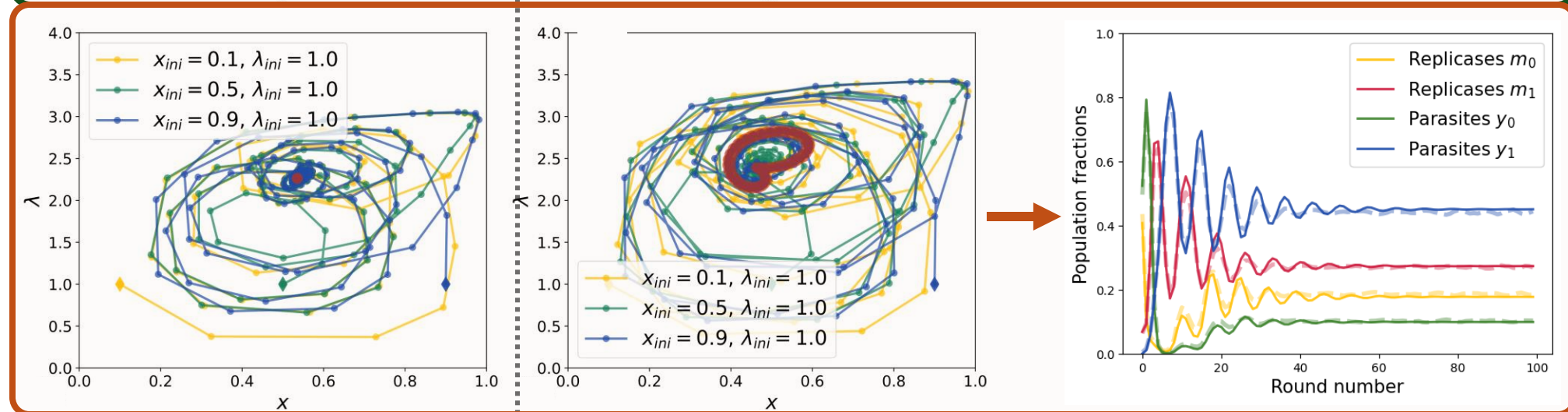
$$\frac{K}{\bar{d}} = 2.9 \quad \text{Carrying capacity} \quad \frac{K}{\bar{d}} = 3.2$$

Dilution

1 parasite  
1 replicase

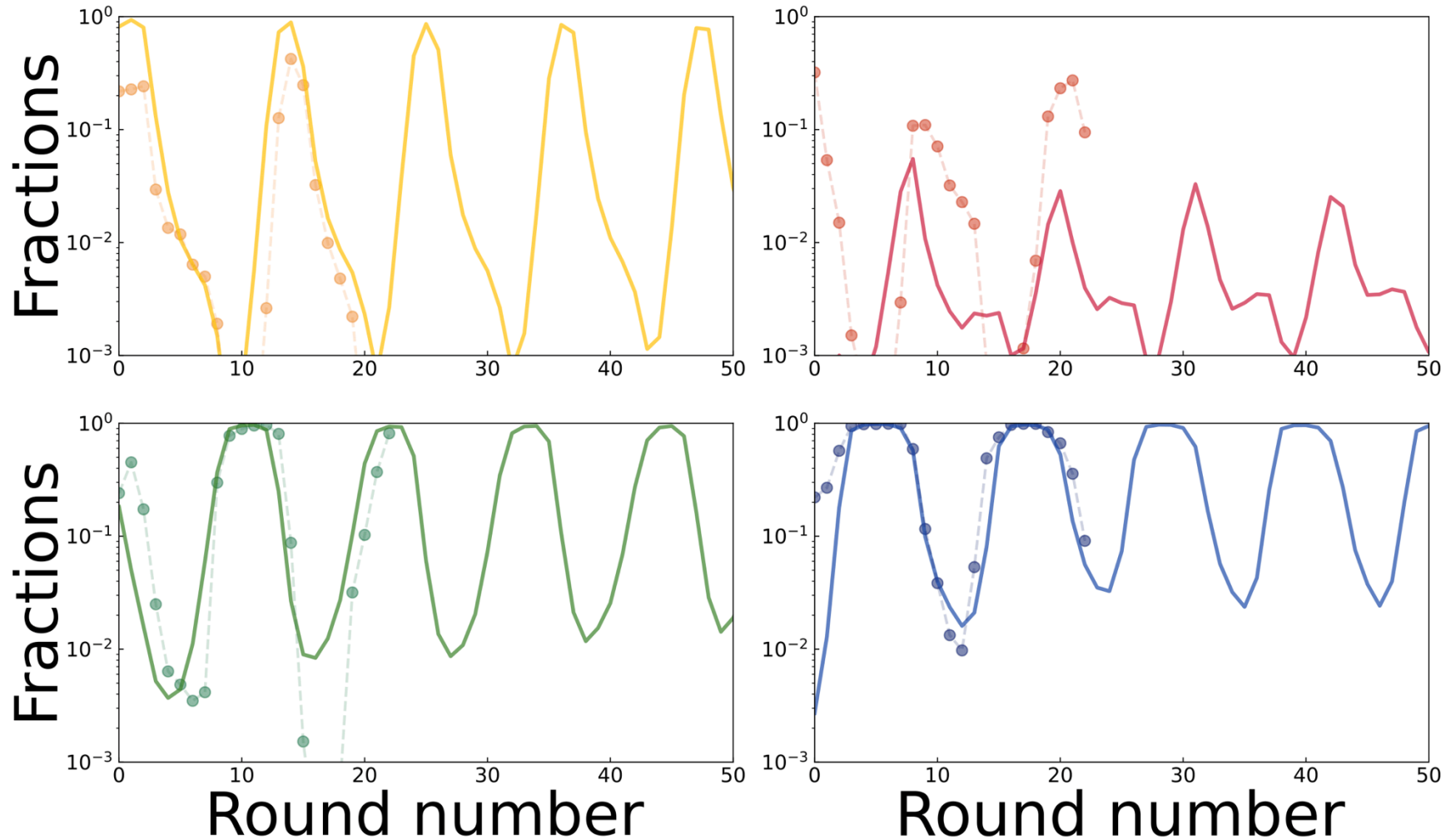


2 parasites  
2 replicases





# Comparison to experiments



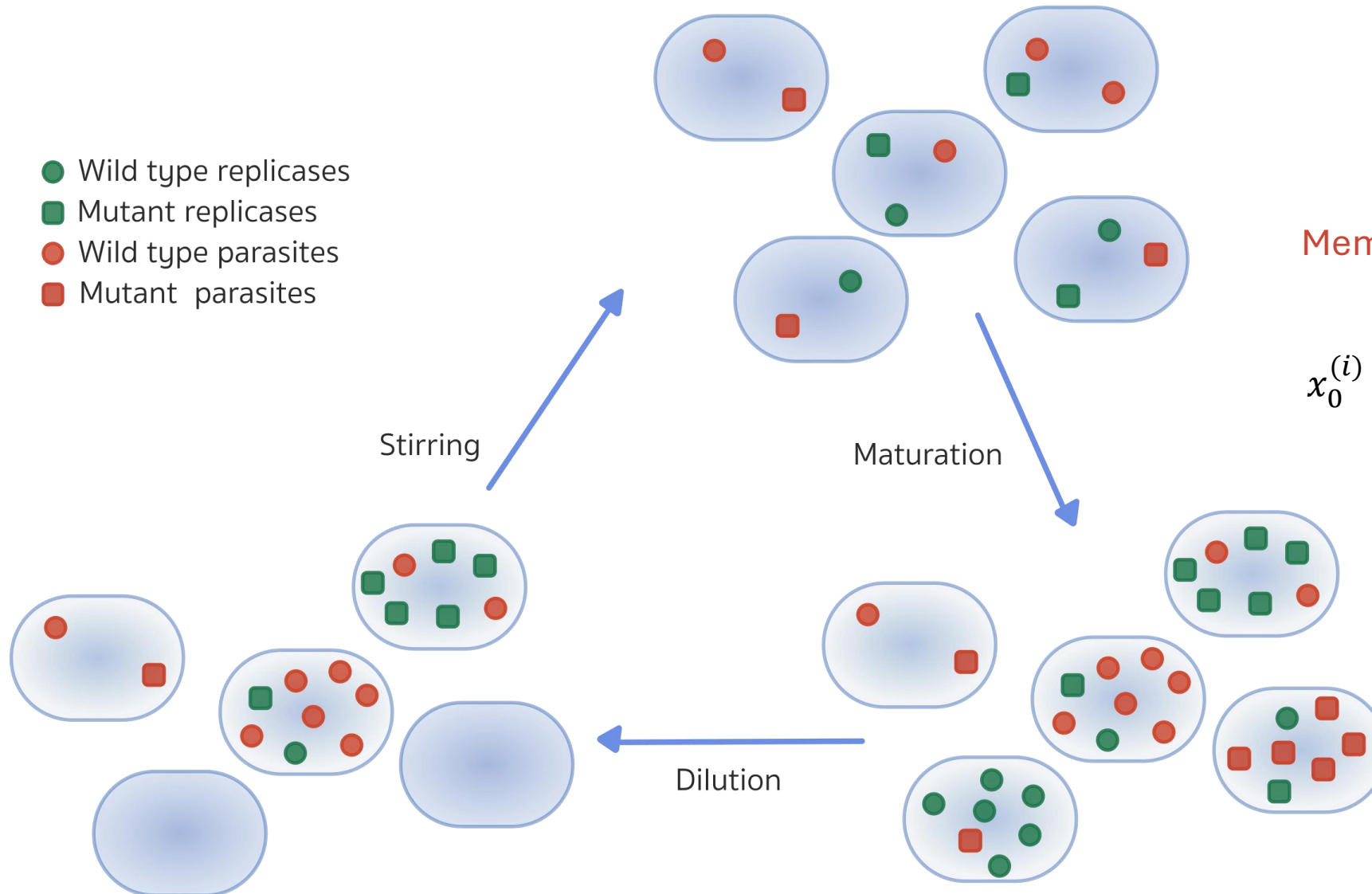
# Stirring

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# Stirring in serial transfer

- Wild type replicases
- Mutant replicases
- Wild type parasites
- Mutant parasites



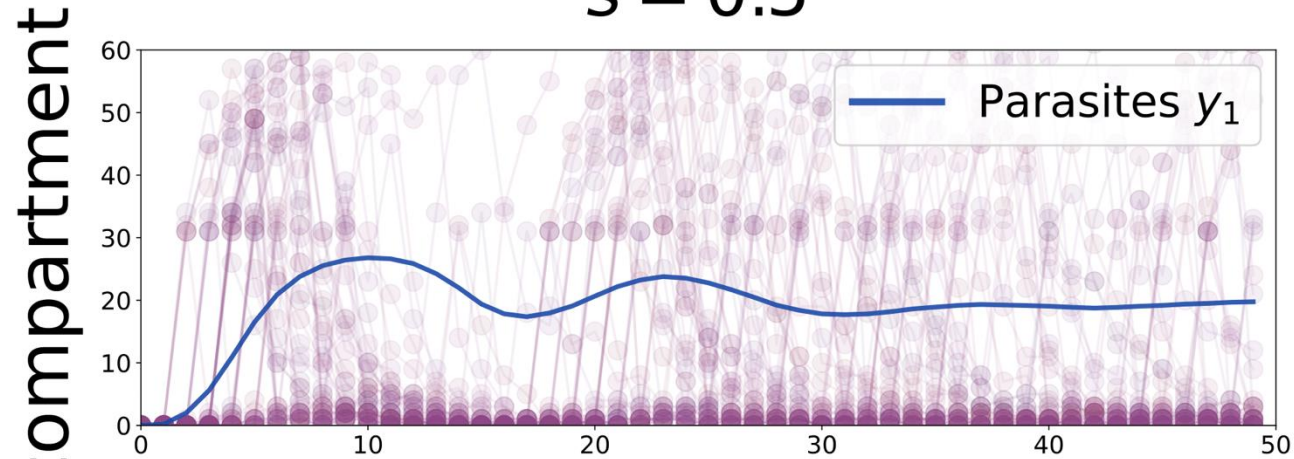
Memory of the compartment

$$x_0^{(i)} = \frac{(1-s)m_0^{(i)}}{(1-s)n^{(i)}} + \frac{\frac{s}{n_{cell}-1} \sum_j m_0^{(j)}}{\frac{s}{n_{cell}-1} \sum_j n^{(j)}}$$

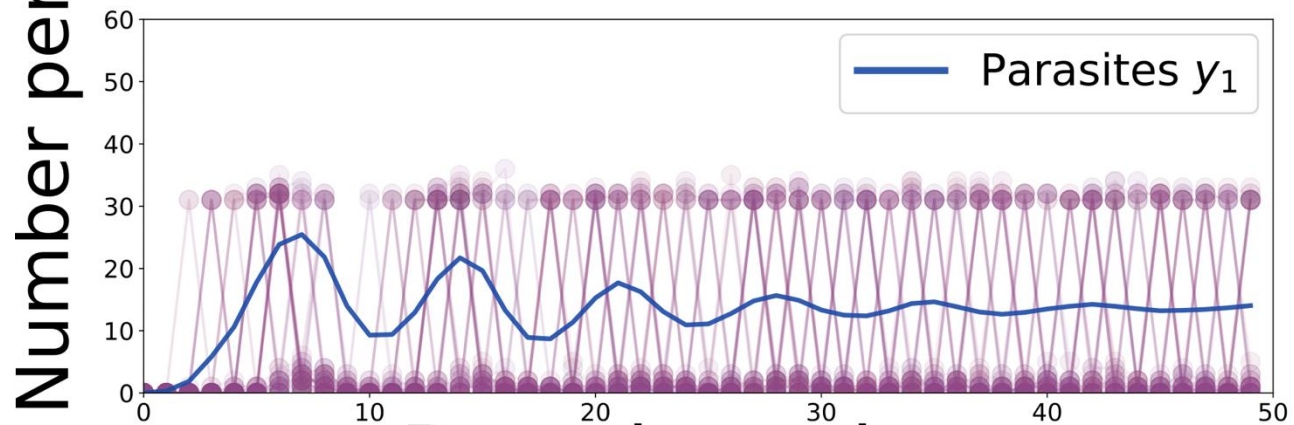
Mean field effect

# Single compartment evolution

$s = 0.5$



$s = 0.99$



Increasing stirring

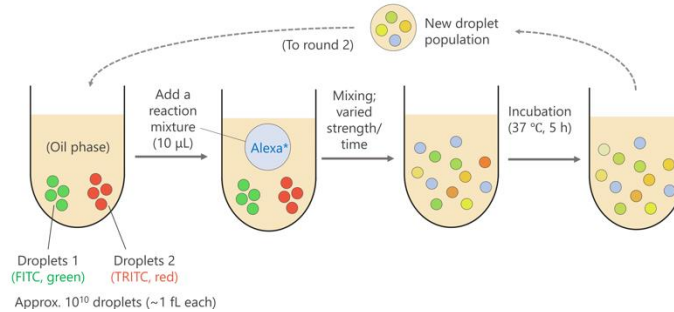


Increasing heterogeneity

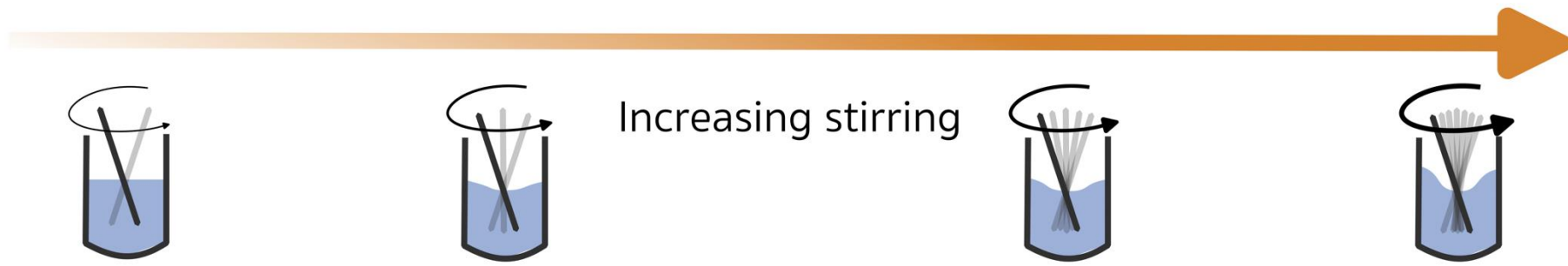
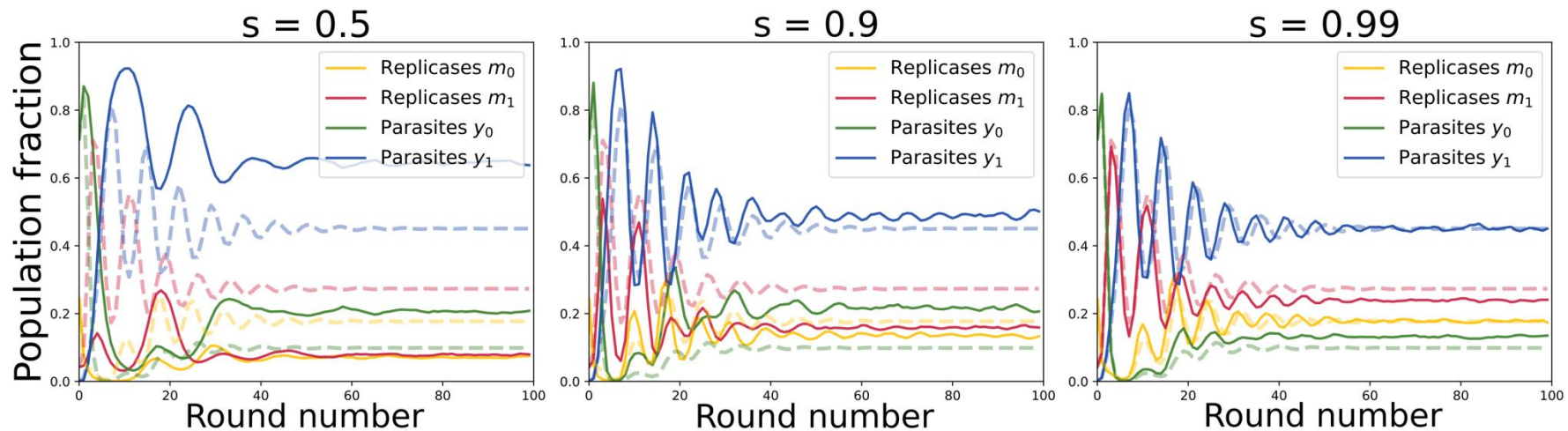
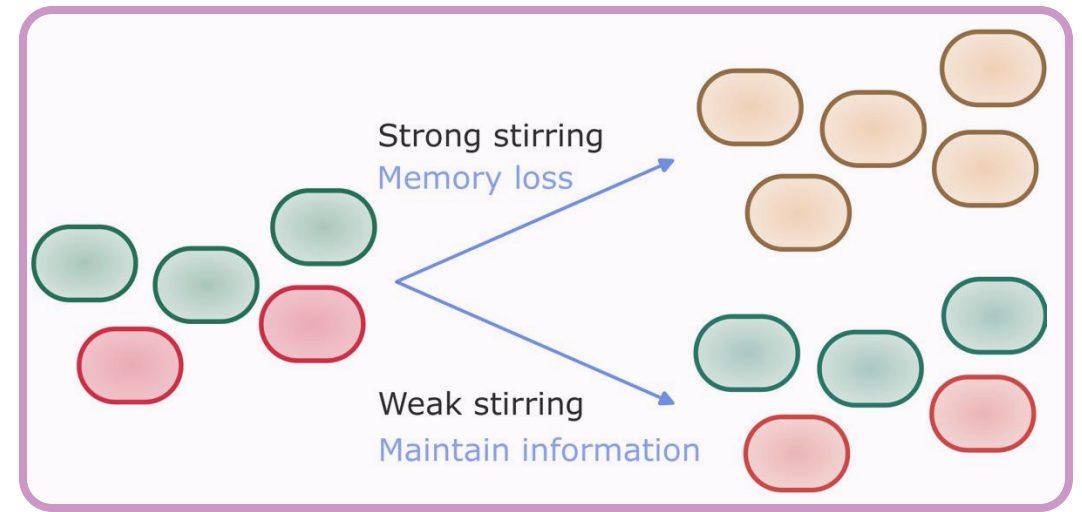


Round number

# Stirring in serial transfer



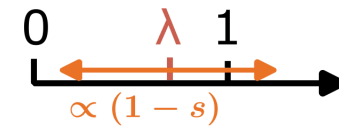
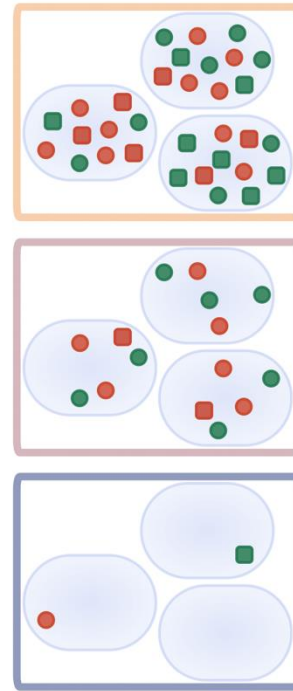
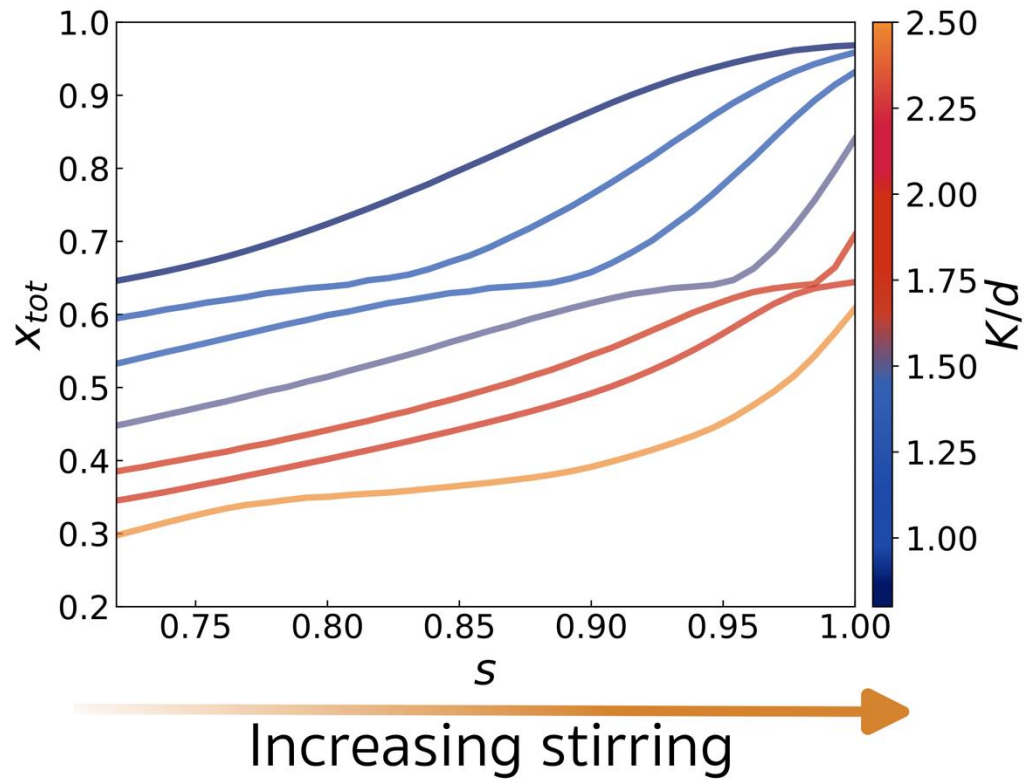
**Tested experimentally**



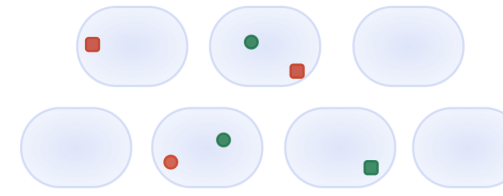


# Stirring favors replicases

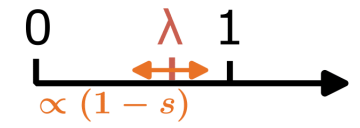
Tested experimentally



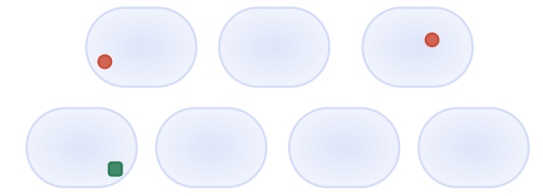
Small  $\lambda$   
High variability



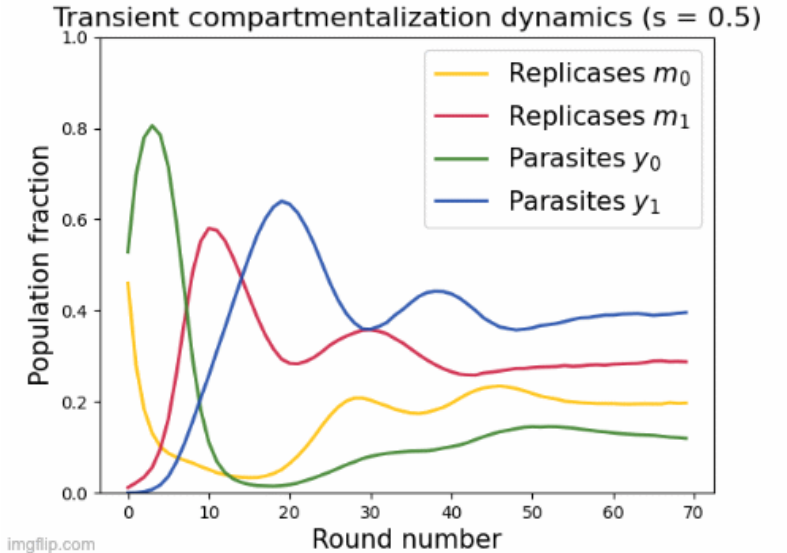
Small  $s$



Small  $\lambda$   
Low variability



Large  $s$





# Conclusion

- **Compartments** along with **stirring** are a way to ensure **coevolution** and appearance of **complexity**.
- More species and strong dilution **stabilizes** the system.
- **Stirring** homogenizes the system, which tend to **isolate parasites** and favors **replicases**.



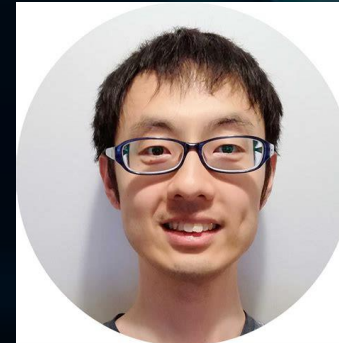
David Lacoste



Ryo Mizuuchi



Ryoka Kuwabara



Taro Furubayashi



Philippe Nghe

# Additional content

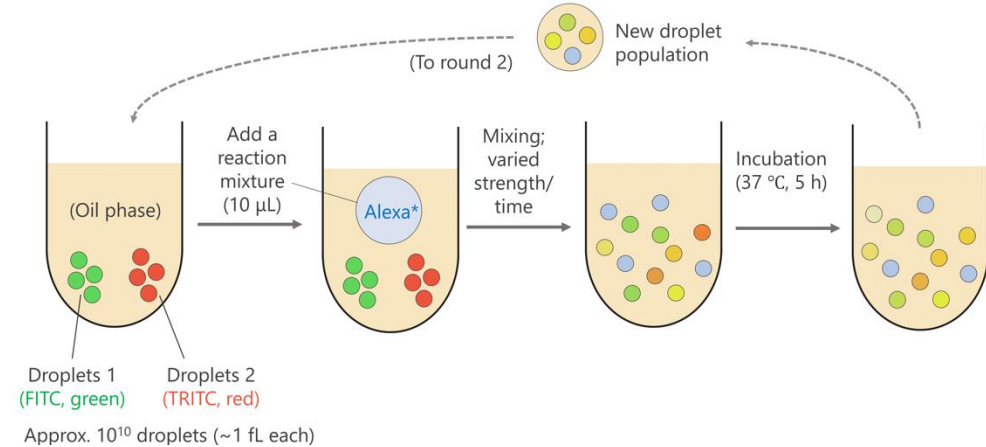
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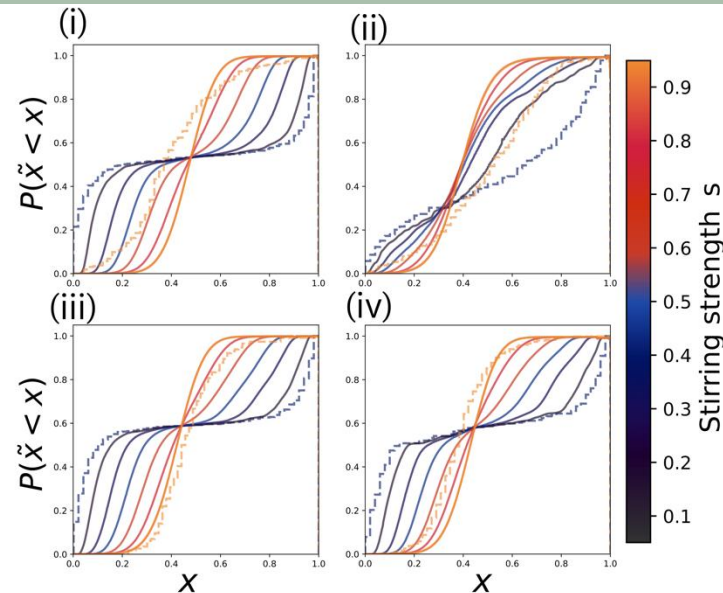
# Experimental tests

With 2 species :

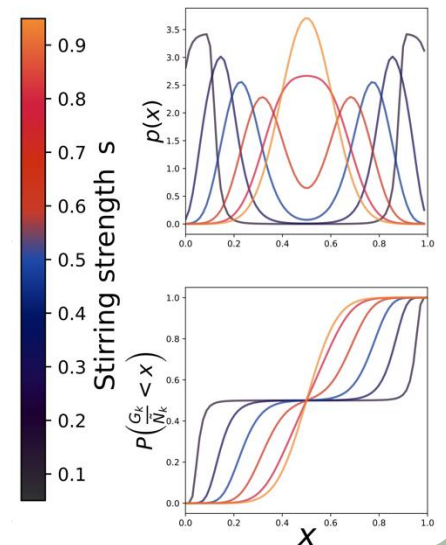
- From bimodal to monomodal distribution
- Depends on stirring process



- (i). Quick, strong stirring
- (ii). Long, weak stirring
- (iii). Medium length, strong stirring
- (iv). Long, strong stirring

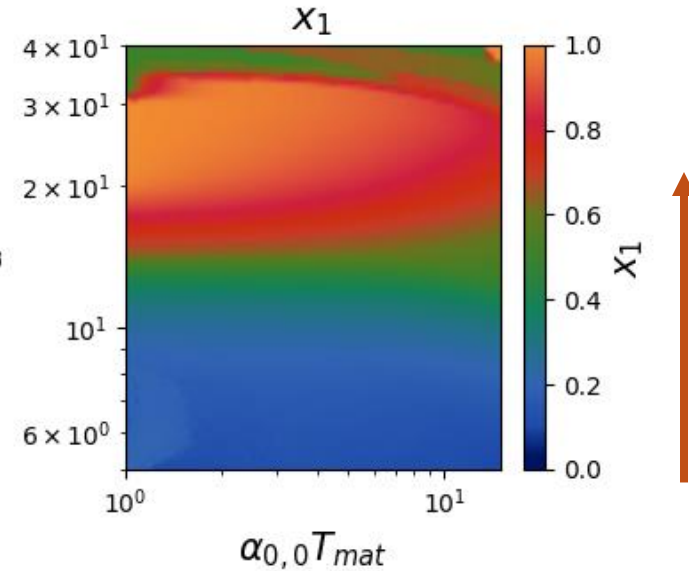
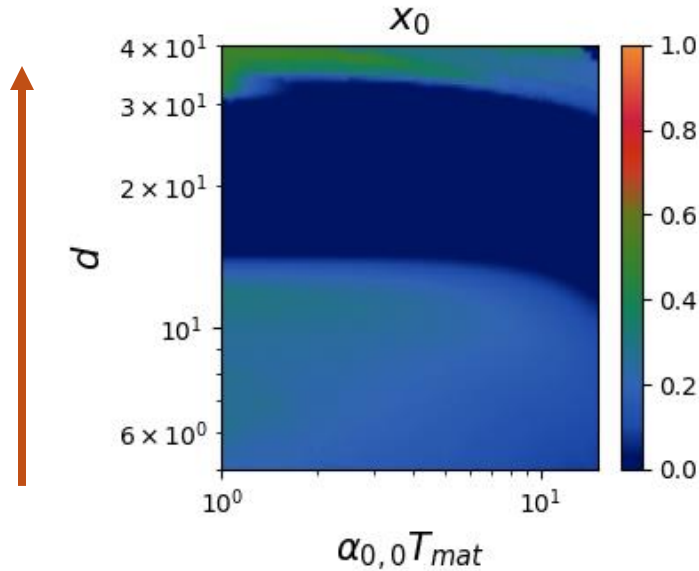


Distribution starting from 2 Dirac peaks.

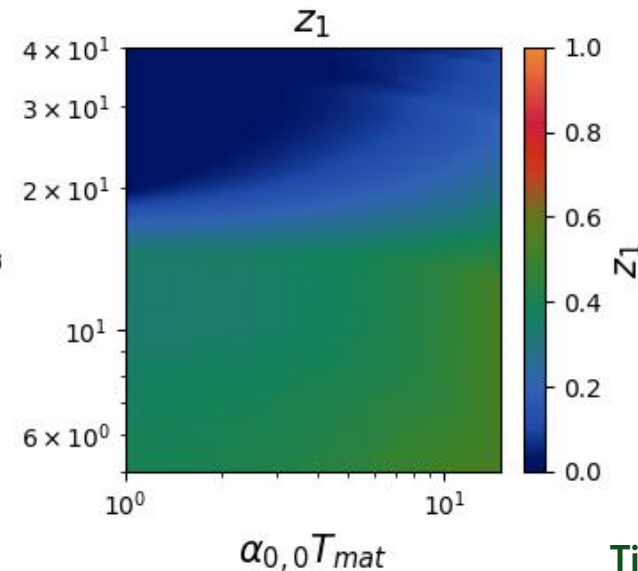
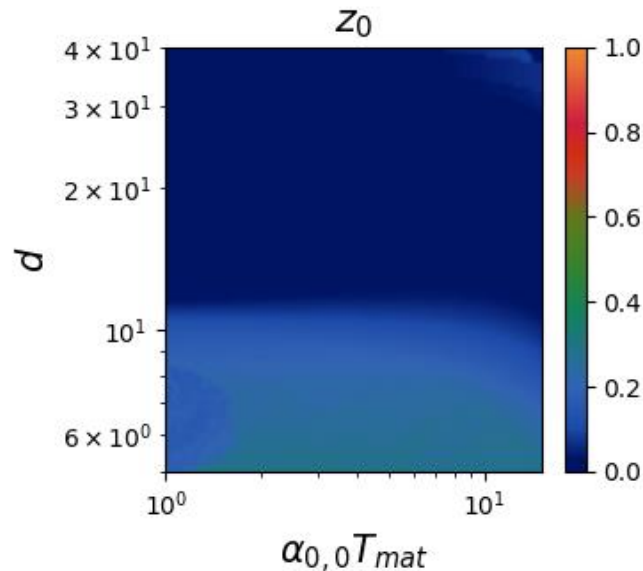


# Influence of dilution factor and maturation time

Very strong dilution:  
 $\lambda \leq 1 \Rightarrow$  No competition



Strong dilution:  
 $\lambda \sim 1 \Rightarrow$  Isolate parasites



Time for maturation:  
Mutations towards parasites