Structured population dynamics with partial sampling

Lab Immersion

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Introduction

Wright-Fisher model

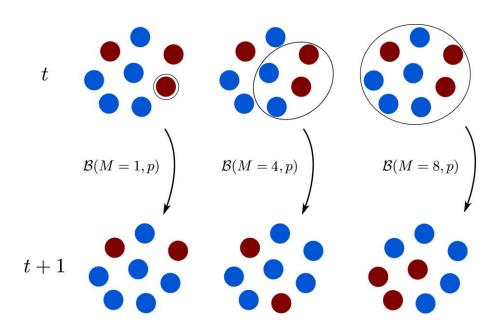
$$P_{i \to j} = \binom{N}{j} p^j (1-p)^{N-j}$$

$$\begin{cases} x = \frac{i}{N} \\ p = \frac{x(1+s)}{1+xs} \end{cases}$$

Moran model

$$P_{i \to j} = \begin{cases} x(1-p) & \text{if } j = i - 1\\ xp + (1-x)(1-p) & \text{if } j = i\\ p(1-x) & \text{if } j = i + 1\\ 0 & \text{otherwise} \end{cases}$$

Introduction: partial sampling



From Alexandre, Arthur, Alia Abbara, Cecilia Fruet, Claude Loverdo, and Anne-Florence Bitbol. "Bridging Wright-Fisher and Moran Models."

Introduction: Fixation probability

N = 1000 $\rho = 1.0$ 10-2 10^{-3} 10^{-3} 10^{-2} 10^{-1} 100 10^{-4}

- Probability to fixate starting from one mutant, given its relative fitness
- Under the diffusion approximation:

$$N \gg 1, s \sim \frac{1}{N}$$

$$\phi(s) = \frac{1 - \exp(-\frac{2}{2-\rho}s)}{1 - \exp(-\frac{2}{2-\rho}Ns)}$$
 with $\rho := \frac{M}{N}$

Methods: Well-mixed case

Simulations:

- Simulate a large number of runs
- Count the number of fixations, and the number of steps before extinction/fixation

Matrix inversions:

• Compute the transition matrix *P*, and its limit by permutating some states

$$P = \begin{bmatrix} 1 & 0 & 0 \\ b_1 & A & b_2 \\ 0 & 0 & 1 \end{bmatrix} \in \mathbb{R}^{(N-1)\times(N-1)} \quad \leftrightarrow \bar{P} = \begin{bmatrix} A & B \\ 0 & I_2 \end{bmatrix}$$

$$\bar{P}^n = \begin{bmatrix} A^n & B_n \\ 0 & I_2 \end{bmatrix} \quad \text{with} \quad B_n = \left(\sum_{k=0}^{n-1} A^k\right) B \underset{n \to \infty}{\longrightarrow} (I_{N-1} - A)^{-1} B \qquad \Longrightarrow \qquad \phi = \left[(I_{N-1} - A)^{-1} B \right]_{0,2}$$

Manhart, Michael, Allan Haldane, and Alexandre V. Morozov. "A Universal Scaling Law Determines Time Reversibility and Steady State of Substitutions under Selection."

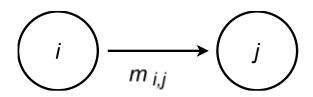
Methods: Structured populations

- Consider a graph of *D* demes, each having *N* individuals
- For each deme, perform a partial sampling step with a weighted average:

$$\tilde{x}^{[i]} = \sum_{k=1}^{D} m_{k,i} x^{[k]}$$

ullet The structure is represented by the migration rate matrix $m=(m_{i,j})_{1\leq i,j\leq D}$

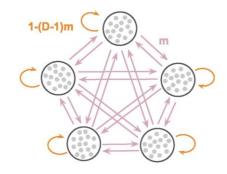
Such that
$$0 \leq m_{i,j} \leq 1$$
, $\sum_{k=1}^{D} m_{k,i} = 1$



Methods: Graphs

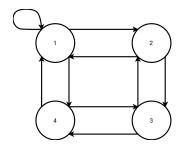
• Clique:

$$m_{i,j} = \begin{cases} 1 - (D-1)m & \text{if } i = j\\ m & \text{otherwise} \end{cases}$$



• Cycle:

$$m_{i,j} = \begin{cases} m & \text{if} \quad j = i - 1 \mod D \\ 1 - (\alpha + 1)m & \text{if} \quad j = i \mod D \\ \alpha m & \text{if} \quad j = i + 1 \mod D \\ 0 & \text{otherwise} \end{cases}$$

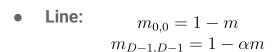


Methods: Graphs

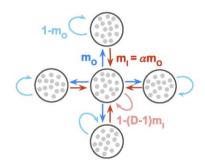
• Star:

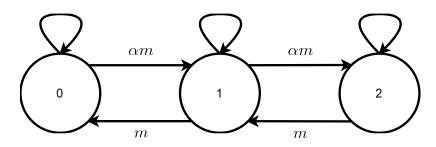
$$\forall i > 0, \quad m_{0,i} = m, \quad m_{i,0} = \alpha m, \quad m_{i,i} = 1 - m,$$

$$m_{0,0} = 1 - (D - 1)\alpha m$$



$$m_{i,i+1} = \alpha m$$
 if $i < D-1$
 $m_{i,i} = 1 - (1+\alpha)m$ if $0 < i < D-1$
 $m_{i,i-1} = m$ if $i > 0$



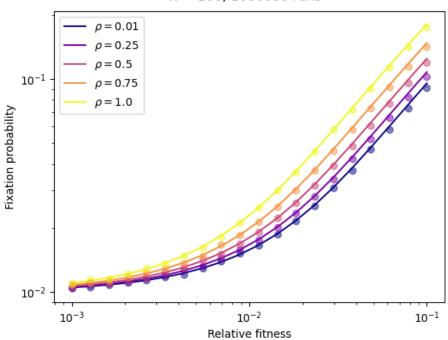


Results: Well-mixed

Well-mixed simulations N = 100, 1000000 runs

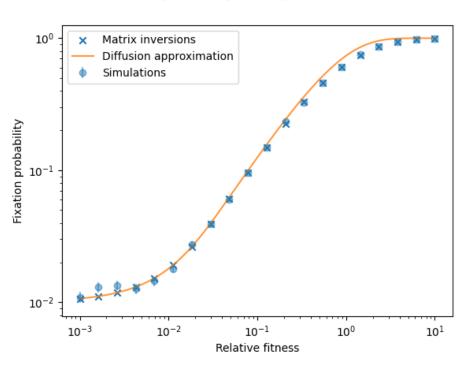
Dots: simulation results

Lines : Diffusion approximation



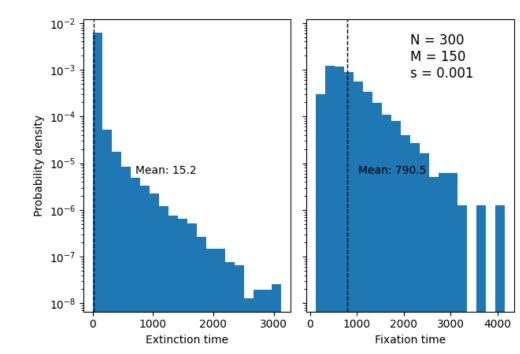
Results: Well-mixed

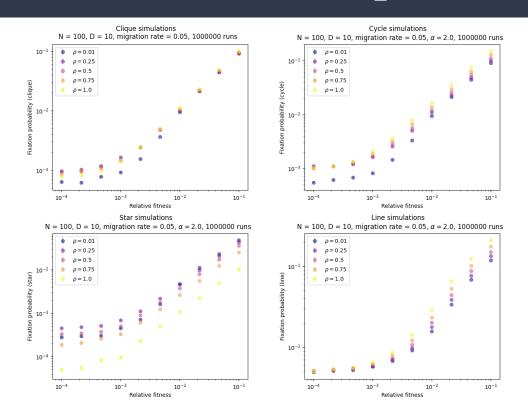
Well-mixed, N = 100, M = 50, 10000 simulations

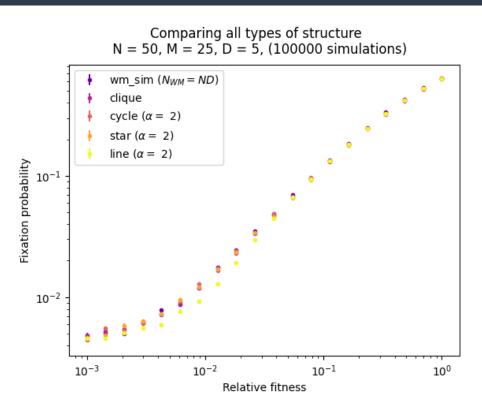


Results: Well-mixed

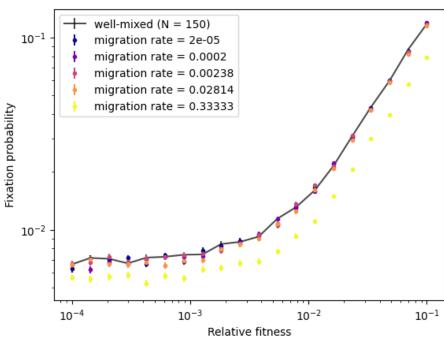
Well-mixed simulations (1000000 runs)

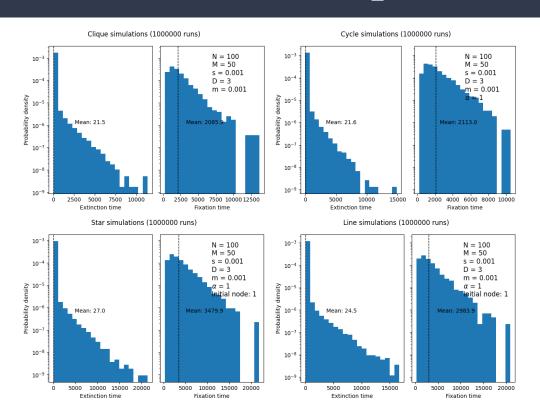




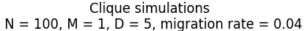


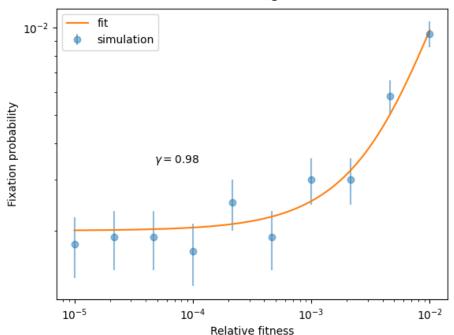
Comparing migration rates on Clique simulations N = 50, M = 25, D = 3, (100000 simulations)





Results: Fitting a model curve



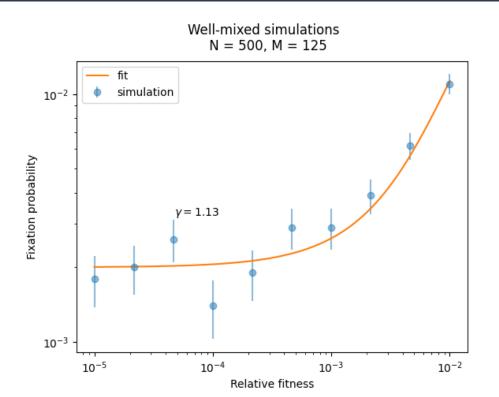


Model curve:

$$\phi_{\gamma}(s) = \frac{1 - \exp(-\gamma s)}{1 - \exp(-\gamma NDs)}$$

(10 000 runs)

Results: Fitting a model curve (WM)



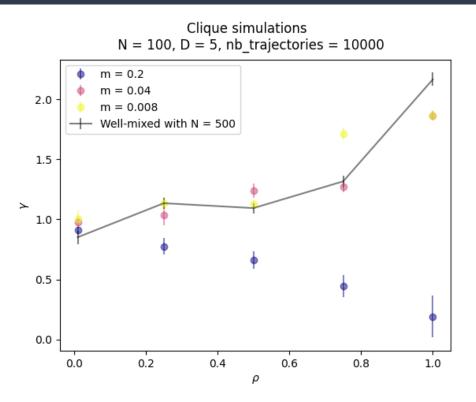
Model curve:

$$\phi_{\gamma}^{WM}(s) = \frac{1 - \exp(-\gamma s)}{1 - \exp(-\gamma N s)}$$

Under the diffusion approximation:

$$\gamma_{th} = \frac{2}{2 - \rho} \approx 1.14 \quad (N = 500, M = 125)$$

Results: Fitting a model curve



Conclusion

- Investigate the results for large migration rates (should behave like in the well-mixed case)
- Alternative ways to compute the weighted fraction:

$$p_{i} = \frac{\tilde{x}^{[i]}(1+s)}{1+\tilde{x}^{[i]} \cdot s} \quad \text{with} \quad \tilde{x}^{[i]} = \sum_{k=1}^{D} m_{k,i} x^{[k]}$$
$$p'_{i} = \sum_{k=1}^{D} m_{k,i} \frac{\tilde{x}^{[k]}(1+s)}{1+\tilde{x}^{[k]} \cdot s}$$

Improve precision (more runs, larger N, D, more data points)

References

- Abbara, Alia, and Anne-Florence Bitbol. "Frequent Asymmetric Migrations Suppress Natural Selection in Spatially Structured Populations." PNAS Nexus 2, no. 11 (November 14, 2023): pgad392. https://doi.org/10.1093/pnasnexus/pgad392.
- Abbara, Alia, Lisa Pagani, Celia García-Pareja, and Anne-Florence Bitbol. "Mutant Fate in Spatially Structured Populations on Graphs: Connecting Models to Experiments." *PLOS Computational Biology* 20, no. 9 (September 6, 2024): e1012424. https://doi.org/10.1371/journal.pcbi.1012424.
- Alexandre, Arthur, Alia Abbara, Cecilia Fruet, Claude Loverdo, and Anne-Florence Bitbol. "Bridging Wright-Fisher and Moran Models." *Journal of Theoretical Biology* 599 (February 21, 2025): 112030.
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- Manhart, Michael, Allan Haldane, and Alexandre V. Morozov. "A Universal Scaling Law Determines Time Reversibility and Steady State of Substitutions under Selection." *Theoretical Population Biology* 82, no. 1 (August 1, 2012): 66–76. https://doi.org/10.1016/j.tpb.2012.03.007.