

Introduction to Computational Biology; An Evolutionary Approach: Genes in Populations: Forward in Time

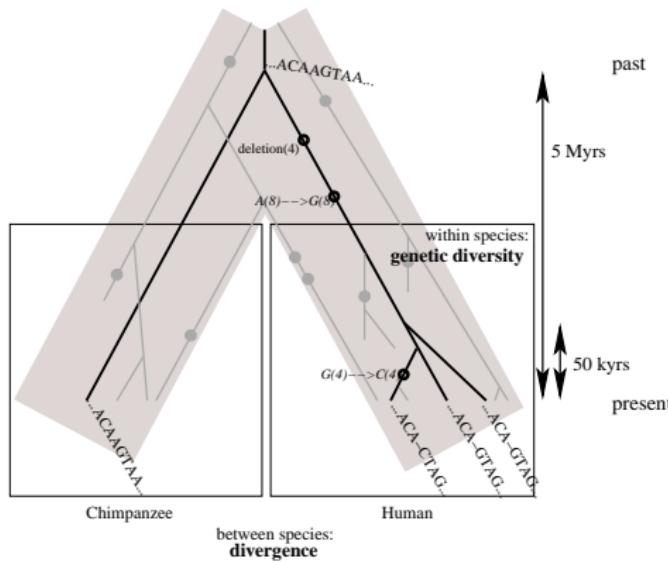
Bernhard Haubold & Thomas Wiehe

Outline

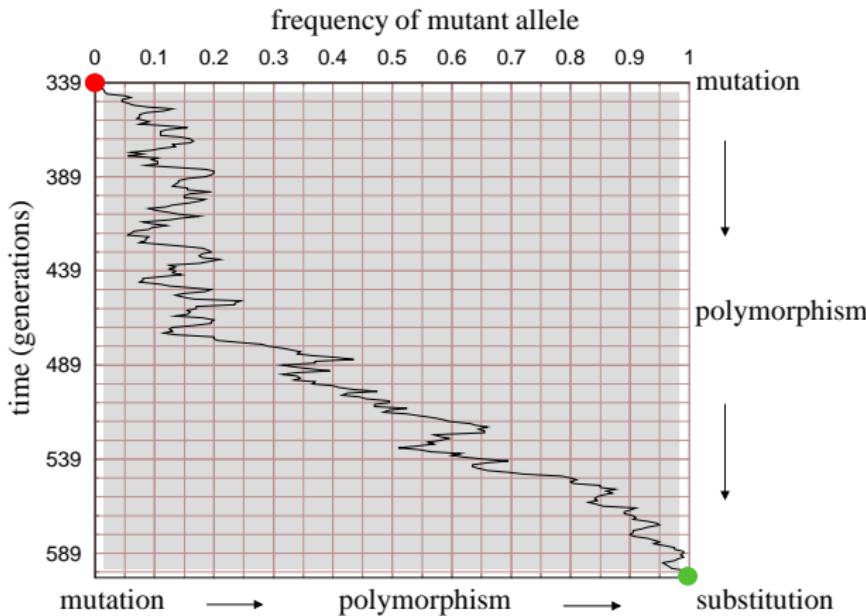
1 Polymorphism & Genetic Diversity

2 Sampling Alleles from Populations

Genealogy of DNA Sequences



Polymorphisms & Substitutions



Types of Polymorphism

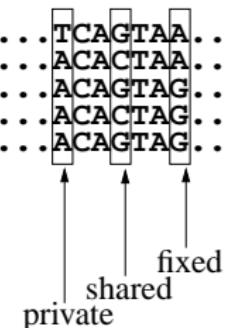
A

single nucleotide polymorphism:

indiv.1 ...ACG**TTCGT**....
 indiv.2 ...AC**A**TTCGT....

B

Chimp1	...TCAG TAA ...
Chimp2	...ACACT AA ...
Human1	...ACAG TAG ...
Human2	...ACACT TAG ...
Human3	...ACAG TAG ...



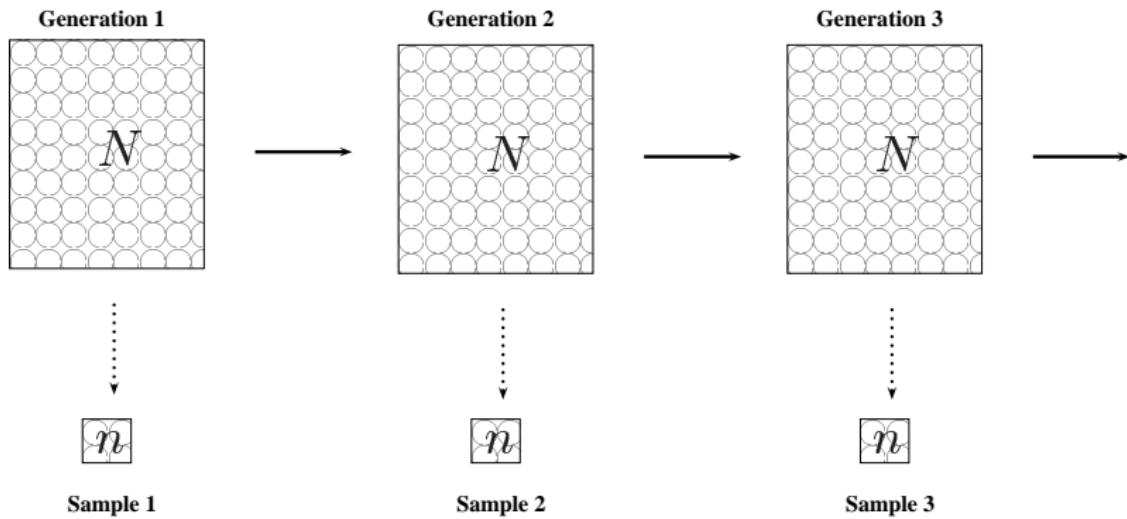
small-tandem-repeat polymorphism:

indiv.1 ...ACGG**GAGAGAGAGA**TTCGT....
 indiv.2 ...ACGG**GAGAGAGA**TTCGT....

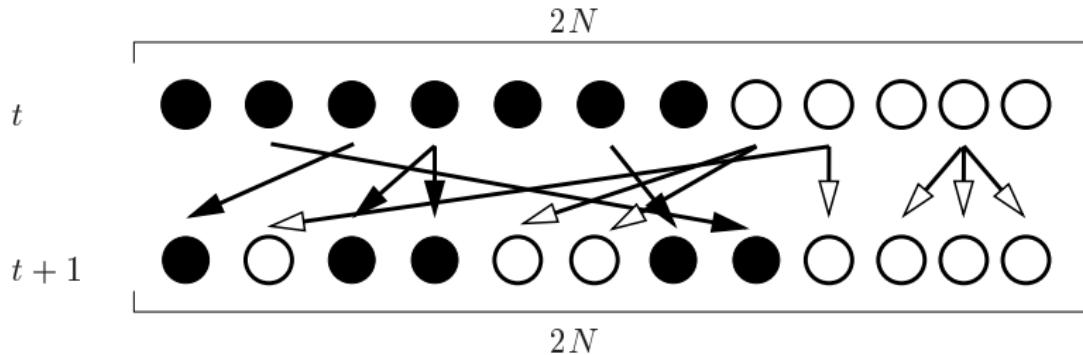
Adh of *D. melanogaster*

Strain	5' Flanking	Exon 1	Intron 1	Exon 2	Intron 2	Exon 3	Intron 3	Exon 4, translated	3' Untranslated	3' Flanking
	CCG	CAATATGGG ∇_1 C ∇_2 GC	T	AC	CCCC	GGAAT	CTCC <u>A</u> CTAG	A ∇_3 C	AGC ∇_4 C ∇_5 T Δ_6	
Wa-SAT.....	.	..	TT.A	CA.TA	AC.....	Δ
Fl-1S	.C	TT.A	CA.TA	AC.....	Δ
Af-SAT ∇ ..1A.	
Fr-S	GTA	.-1.	TA.
Fl-2S	...	AG...A.TC...A	G	GT	c 3.	
Ja-S	.C	GT.T.CA	c 4.	...T	...
Fl-F	.C	GGTCTCC.	c 4.	
Fr-F	TGC	AG...A.TC ∇ G ∇ ..	GGTCTCC.	c 4 G	
Wa-F	TGC	AG...A.TC ∇ G ∇ ..	GGTCTCC.	c 4 G	
Af-F	TGC	AG...A.TC ∇ G ∇ ..	GGTCTCC.	c 5 G	
Ja-F	TGC	AGGGGA... ∇ ..T.	GA.	..G..	..GTCTCC.	c 4.	-1..
Polym. Sites	3	0	12	1	2	4	5	9	2	5
Nucleotides ¹	63	87	690	99	65	405	70	264	178	767
% polymorphic	4.7	0	1.7	1.0	3.1	1.0	7.1	3.5	1.1	0.6

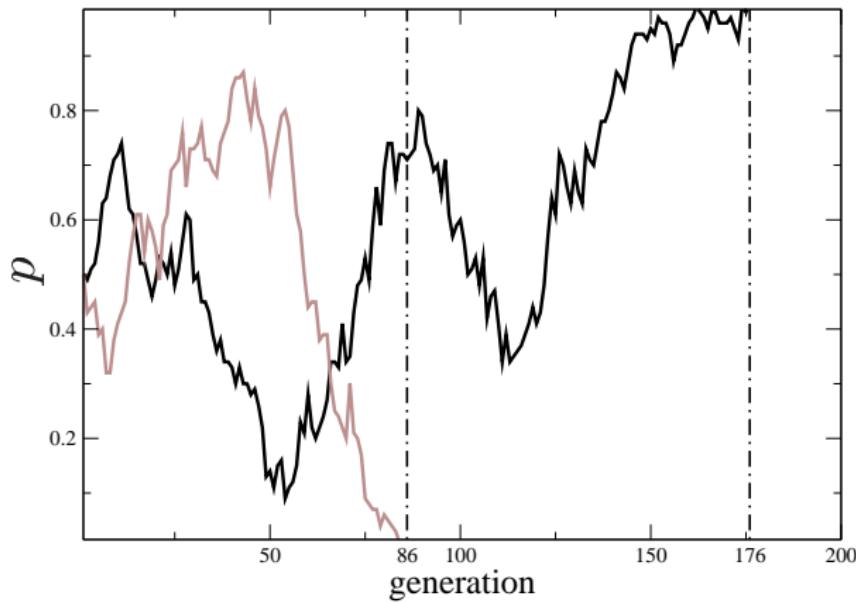
Evolution Forward in Time



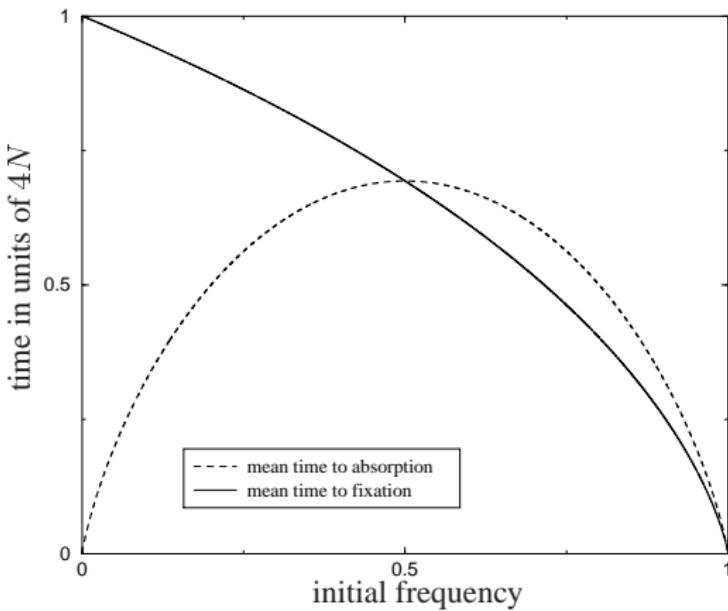
Evolution in two Generations



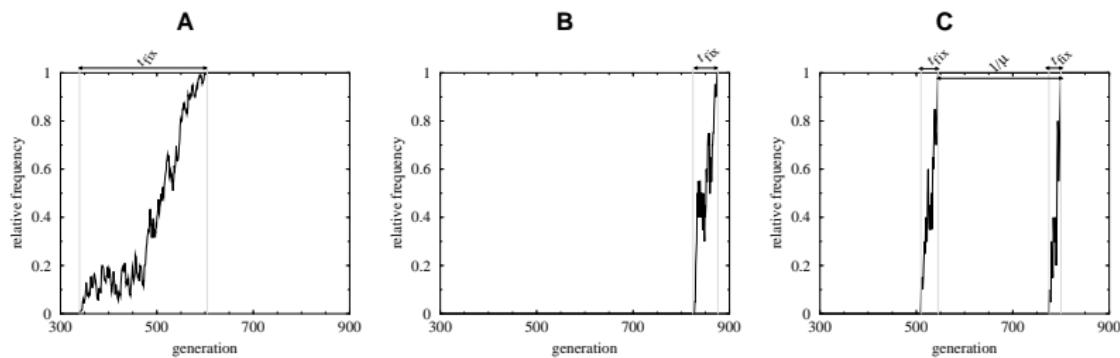
Evolution under Drift



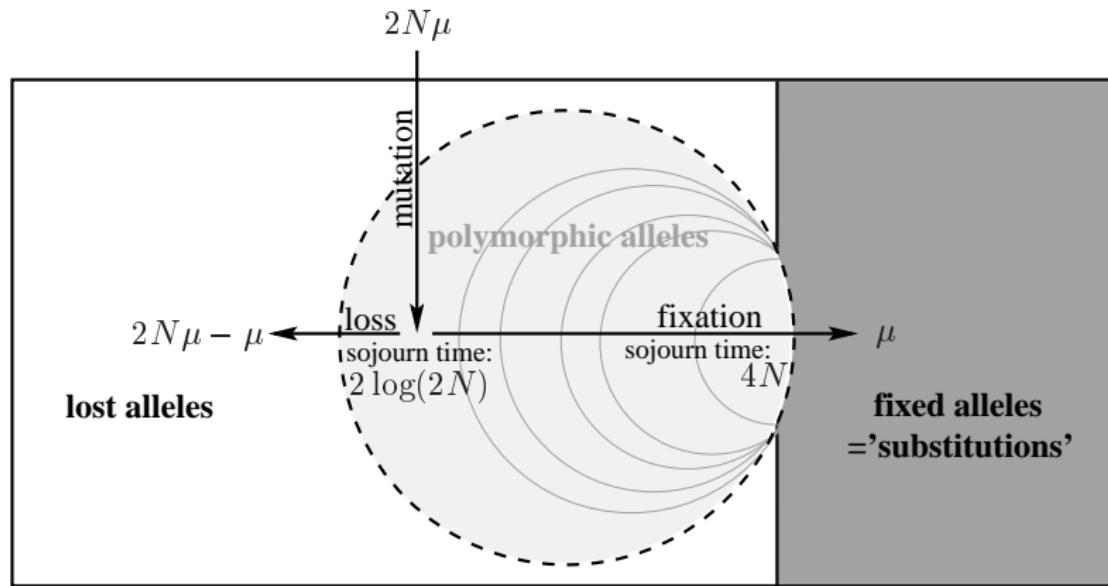
Absorption & Fixation



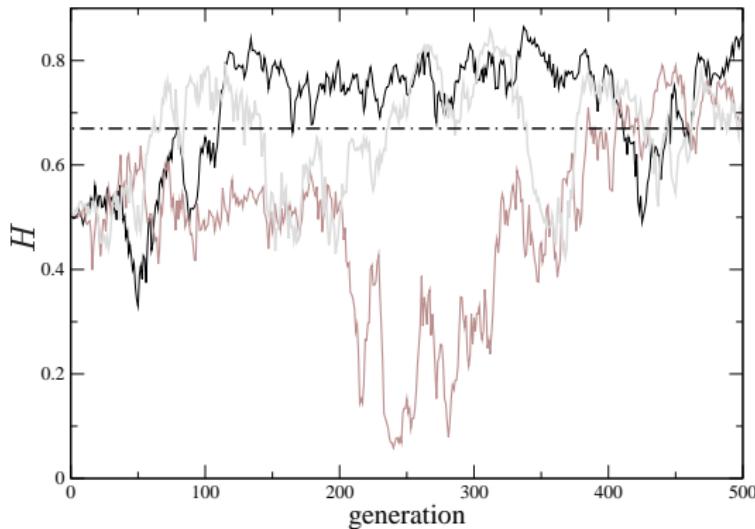
Mutation Drift Balance



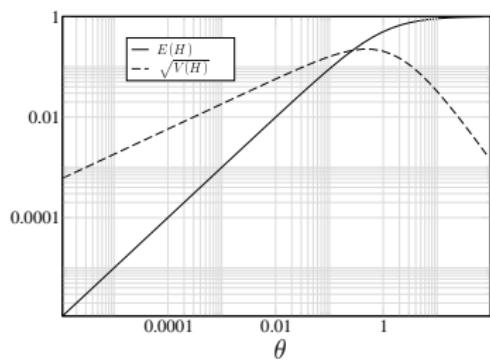
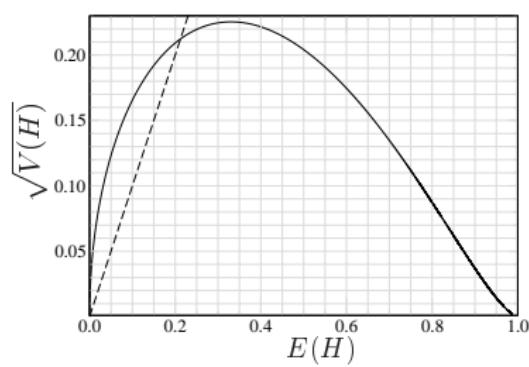
Mutation Drift Balance



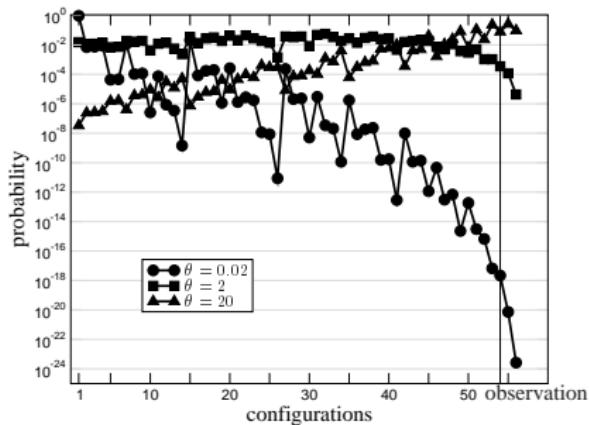
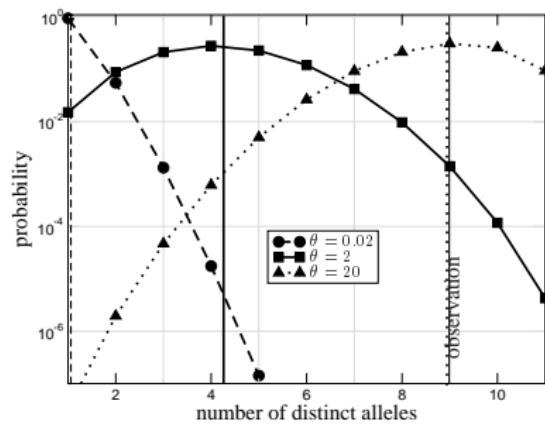
Simulation of Heterozygosity



Heterozygosity and Mutation Rate

A**B**

Sample Configurations

A**B**

Expectations

θ	$\underline{\kappa}^\dagger$	$E(\mathcal{K}_{11})$	$V(\mathcal{K}_{11})$	S^\ddagger	$E(C_1(11))$
0.02	1.020	1.058	0.057	0.586	0.022
2.00	3.000	4.206	1.947	5.858	1.833
6.12	7.120	6.629	2.234	17.925	4.176
20.0	21.00	8.945	1.550	58.579	7.333