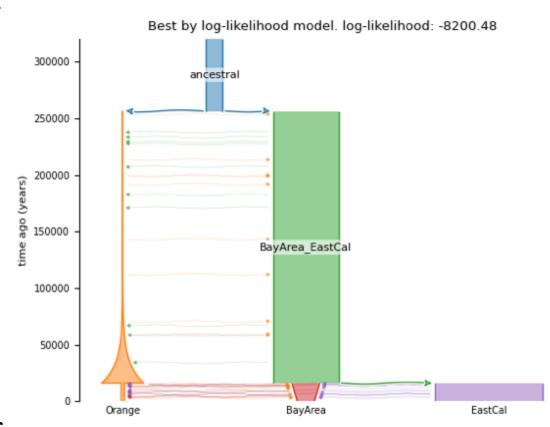
### Part III: Demographic analysis

• Demographic history includes the history of divergence, fluctuations in effective population size, and gene flow among populations.

 Along with population structure certain demographic histories can result in false positive outliers in selection scans.

• Demographic inference central to studies of biogeography, speciation, conservation, etc.



### Demographic methods

• Genealogy samplers (e.g. IMa, Lamarc, Migrate) – Kuhner 2009

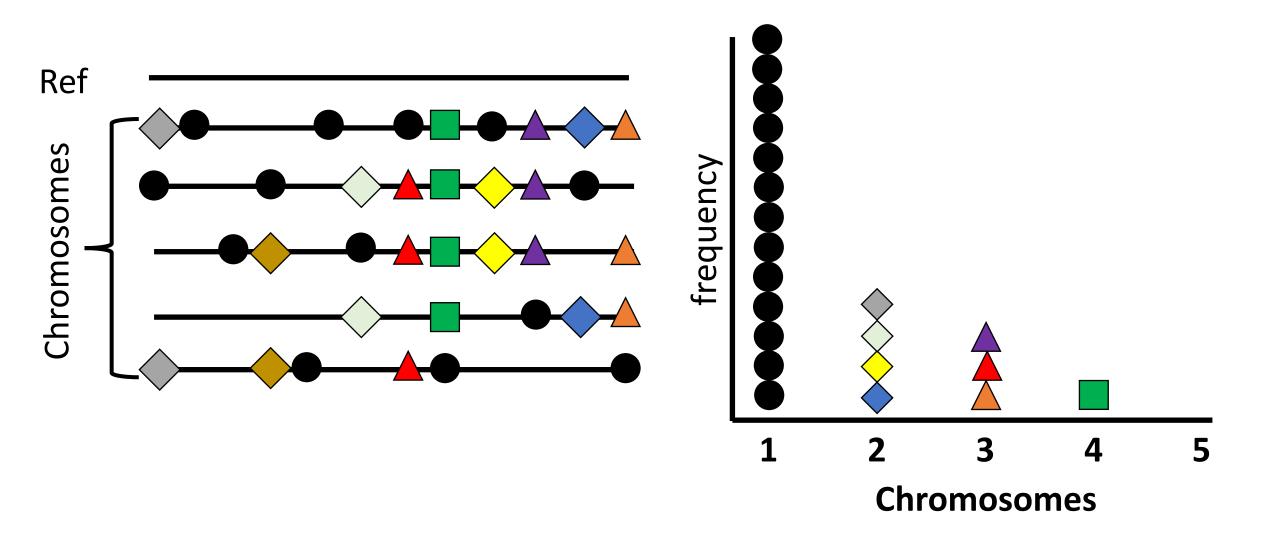
G-PhoCS (Gronau et al. 2011)

Approximate Bayesian computation (ABC)

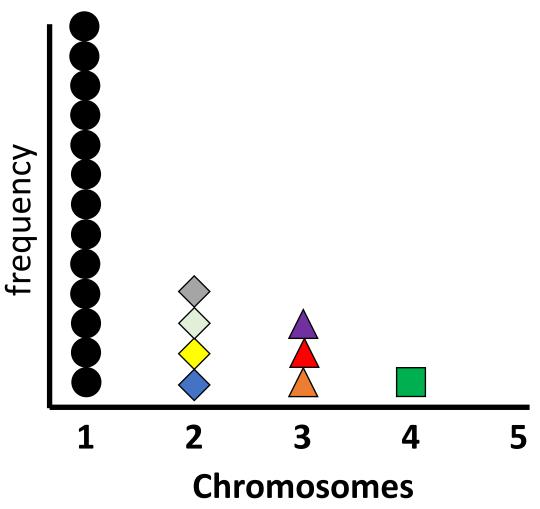
Sequentially Markovian Coalescent (e.g. SMC+, PSMC) -- Spence et al.
2018

• Site frequency spectrum based inference (e.g. ∂aði, fastsimcoal2)

### The site frequency spectrum (1D)



## Many pop. gen. statistics are summaries of the SFS

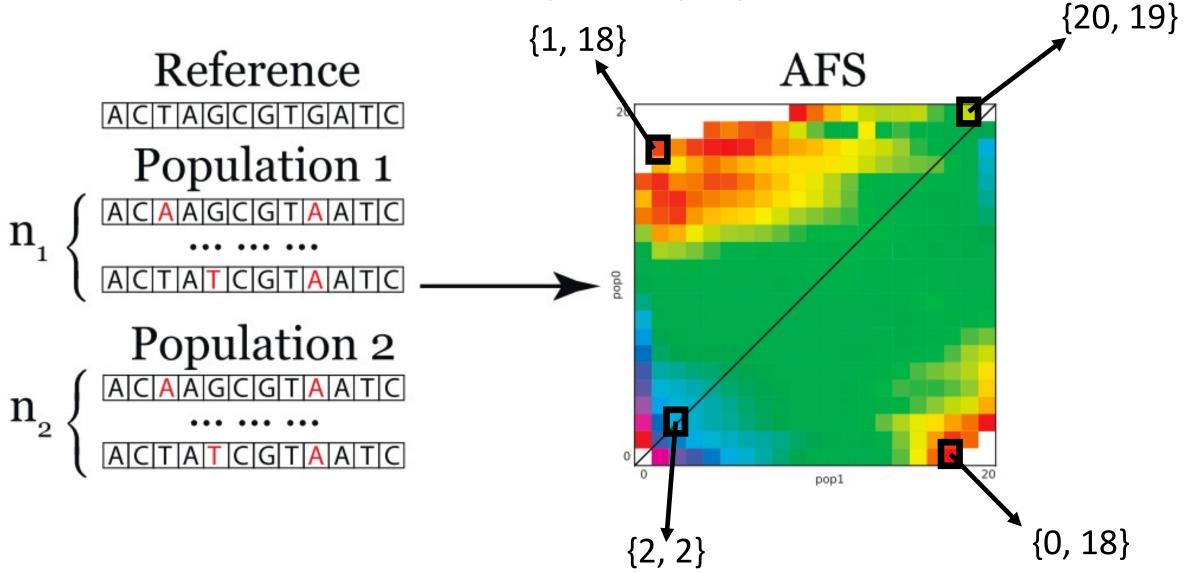


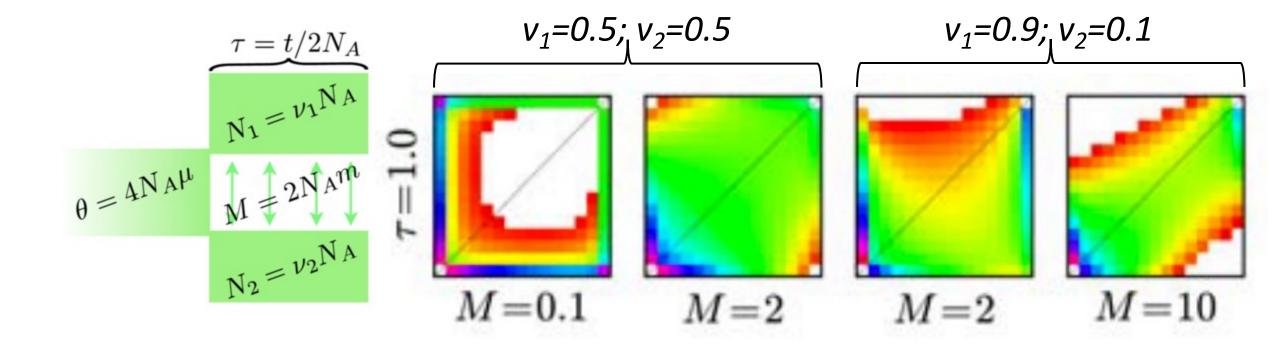
Segregating sites

**Nucleotide diversity** 

Tajima's D

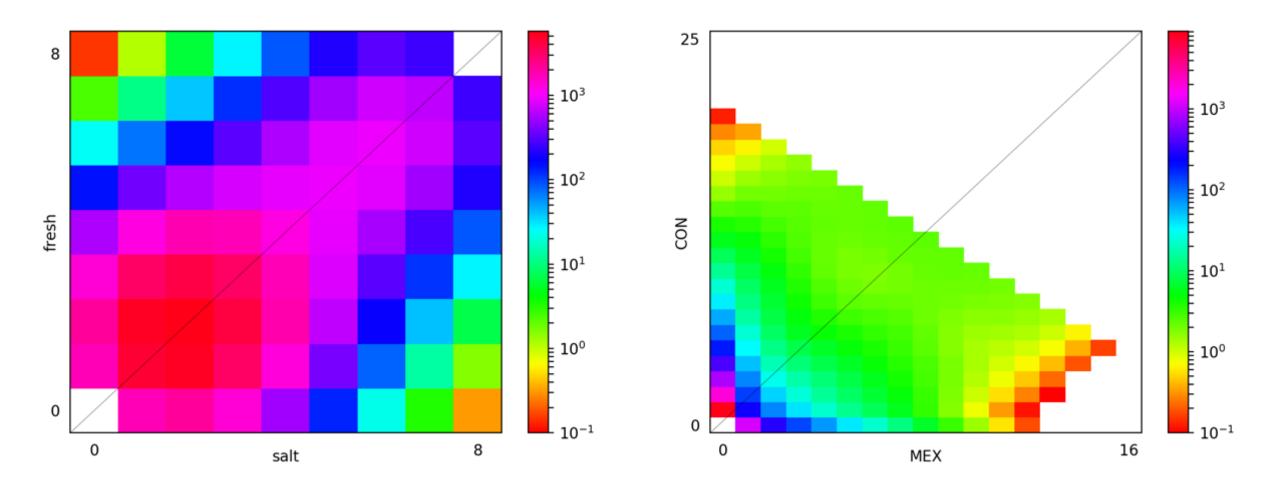
Joint Site Frequency spectrum (2D)





Gutenkunst et al. 2009 PLoS Genet.

### Polarized vs. folded site frequency spectrum



# Approaches to fitting demographic models to the site frequency spectrum.

∂a∂i – diffusion approximation (Gutenkunst et al. 2009).

3-5 populations

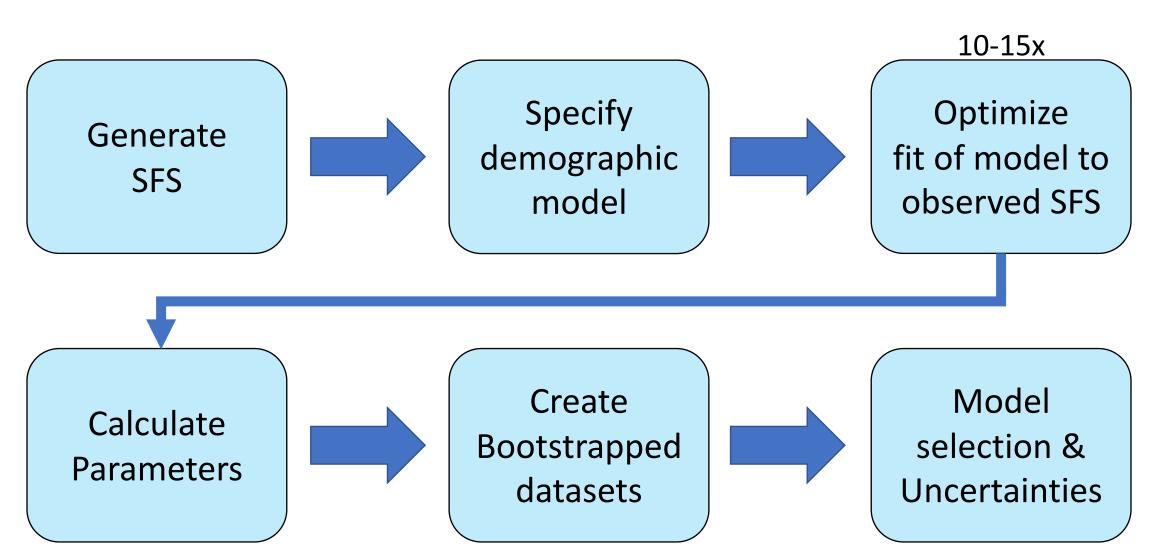
• Moments (Jouganous et al. 2017).

• Fastsimcoal2 – Coalescent based approach (Excoffier et al. 2013).

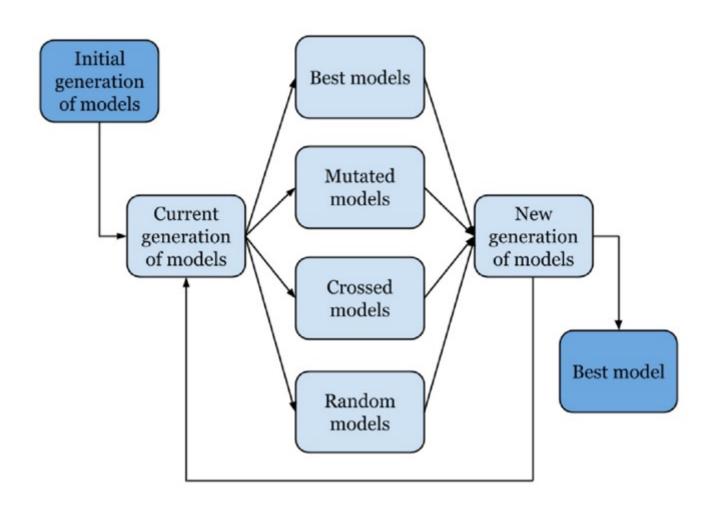
• Momi – (Kamm et al. 2017).

Unlimited populations

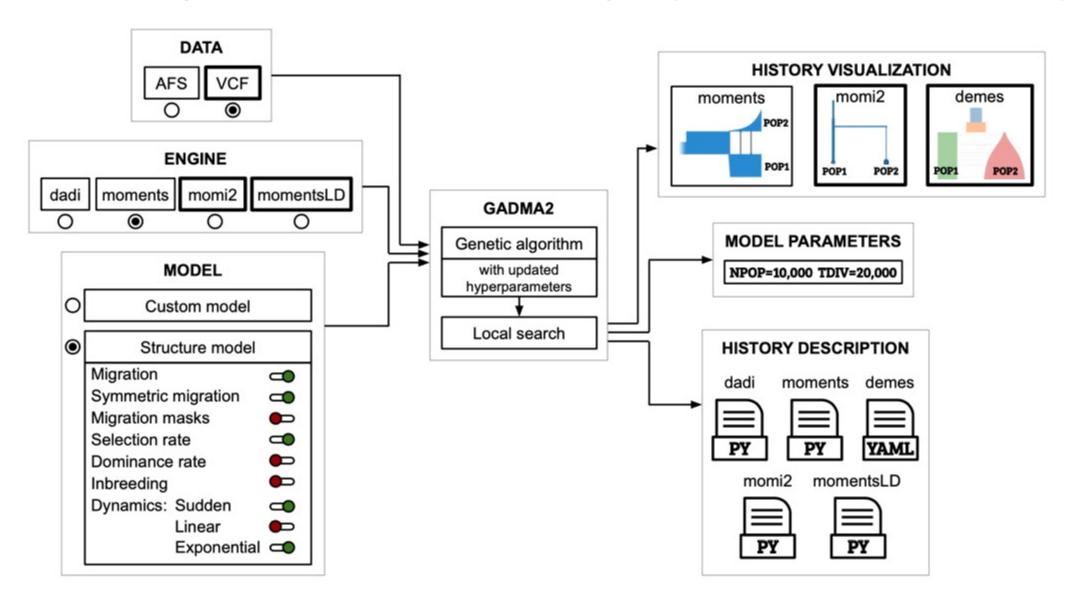
## Pipeline for demographic inference with dadi and moments



### Genetic Algorithm for Demographic Model Analysis



### Genetic Algorithm for Demographic Model Analysis



#### Additional considerations.

- All demographic models are wrong!
- Can be very sensitive to messy data, leads to model misspecification, stacking up against parameter boundaries (e.g., Rosen et al. 2018).
- Bottlenecks can be very challenging to accurately infer. E.g. many datasets underpowered to infer ancient bottlenecks, long versus severe bottlenecks can be conflated (e.g. Terhorst & Song 2015).
- Linked selection causes bias (e.g. smaller population sizes, more recent divergence), non-genic regions preferred for neutral demographic inference (e.g. Schrider et al. 2016).
- Number of individuals: some demographic histories require few individuals (e.g. 5-10) whereas others can require dozens (See Robinson et al. 2014 for some guidelines).





#### **METHODS IN POPULATION GENOMICS**

A website listing current methods in population genomics

https://methodspopgen.com



**HADDOCK • DUNN**