Genetics and population analysis

POPdemog: visualizing population demographic history from simulation scripts

Ying Zhou^{1,*}, Xiaowen Tian¹, Brian L. Browning^{1,2} and Sharon R. Browning¹

¹Department of Biostatistics and ²Division of Medical Genetics, Department of Medicine, University of Washington, WA 98195, USA

*To whom correspondence should be addressed. Associate Editor: Oliver Stegle

Received on September 22, 2017; revised on December 11, 2017; editorial decision on March 22, 2018; accepted on March 23, 2018

Abstract

Summary: We present POPdemog, an R package which converts coalescent simulation program input parameters into a visual representation of the demographic model. This package is useful for preparing figures, for checking that demographic simulation parameters have been correctly specified, and for understanding demographic models that other researchers have used to simulate genetic data. The POPdemog package supports the ms, msa, msHot, MaCS, msprime, scrm and Cosi2 programs, and includes options for customizing the output figures.

Availability and implementation: The POPdemog package and its tutorial can be freely down-loaded from https://github.com/YingZhou001/POPdemog.

Contact: yz001@uw.edu

Supplementary information: Supplementary data are available at *Bioinformatics* online.

1 Introduction

Many simulation tools have been developed to support population and evolutionary genetic studies, and many of these programs are able to generate genetic data under complex demographic models (Hoban *et al.*, 2012). Often the demographic models include several populations with time-varying population sizes and multiple migration events. In this case, the large number of demographic parameters makes it difficult to understand the demographic model directly from the simulation program command line (see Box 1).

We have developed the POPdemog R package to capture the complex demographic information from simulation scripts or parameter files, and output the demographic model as a figure (Fig. 1). The main output figure shows the changes in population sizes and migrations events. When multiple migrations happen at a particular time, additional figures can be generated to give fine-scale representation of the overlapping migration events. Currently, POPdemog supports the input to the ms and msa (Hudson, 2002), msHot (Hellenthal and Stephens, 2007), MaCS (Chen *et al.*, 2009), scrm (Staab *et al.*, 2015), msprime (Kelleher *et al.*, 2016) and Cosi2 (Shlyakhter *et al.*, 2014) programs. Box 1. MaCS command (Vernot *et al.*, 2016) for modified Tennessen model (Tennessen *et al.*, 2012):

macs 2025 1500000 -i 10 -r 3.0e-04 -t 0.00069 -T -I 4 10 1006 1008 1 0 -n 4 0.205 -n 1 58.00274 -n 2 70.041 -n 3 187.55 -eg 0.9e-10 1 482.46 -eg 1.0e-10 2 570.18 -eg 1.1e-10 3 720.23 -em 1.2e-10 1 2 0.731 -em 1.3e-10 2 1 0.731 -em 1.4e-10 3 1 0.2281 -em 1.5e-10 1 3 0.2281 -em 1.6e-10 2 3 0.9094 -em 1.7e-10 3 2 0.9094 -eg 0.007 1 0 -en 0.007001 1 1.98 -eg 0.007002 2 89.7668 -eg 0.007003 3 113.3896 -eG 0.031456 0 -en 0.031457 2 0.1412 -en 0.031458 3 0.07579 -eM 0.031459 0 -ej 0.03146 3 2 -en 0.0314601 2 0.2546 -em 0.0314602 2 1 4.386 -em 0.0314603 1 2 4.386 -eM 0.0697669 0 -ej 0.069767 2 1 -en 0.0697671 1 1.98 -en 0.2025 1 1 -ej 0.9575923 4 1 -em 0.06765 2 4 32 -em 0.06840 2 4 0

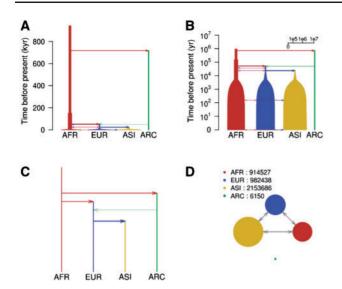


Fig. 1. Visual representations of demography in MaCS simulation script for a modified Tennessen model (Box 1) (Vernot *et al.*, 2016). AFR: Africans; EUR: Europeans; ASI: Asians; ARC: Archaic humans. (**A**) Use a time unit of 1000 years and scale the population size logarithmically. (**B**) Zoom in on the recent events and scale the population size logarithmically. (**C**) 'topology' of migrations with rates \geq 0.001 per generation. (**D**) Migrations and population sizes 1000 years ago, with population sizes scaled linearly. A–C are generated by the *PlotMS()* function, **D** is generated by the *PlotMig()* function

 Table 1. Options for the parameters controlling scaling of population size and event time

size.scale	ʻlog'	Scale the population size logarithmically, with the base parameter log.base [default]
	'linear'	Scale population size linearly, with scale parameter linear.scale
	'topology'	Ignore population size and event times, output only the topological relationship
time.scale	'4Ne'	Set time unit to 4Ne [default]
	'kyear'	Set time unit to 1000 years
	'generation'	Set time unit to generations
	'years'	Set time unit to years
	'log10year'	Scale time with function
		'log10(years before present)'

Note: Further details can be found in the program documentation and the tutorial file.

2 Input and output

The input to our program is the command line or parameter file for invoking the simulation program, the simulation program name, and the baseline effective population size. Many coalescent-based simulation programs allow for arbitrary scaling of simulation parameters relative to an unspecified baseline effective population size; thus the effective population size is required by our program in order to adjust the scale. We also provide options to customize the appearance of the output figure such as the time scale, branch positions, branch colors and branch widths for each population. Options for controlling scaling of time and population size are described in Table 1. For example, setting size.scale = 'log' shows population sizes on a log scale, which is helpful when population size has undergone exponential growth (Fig. 1A), and setting time.scale = 'log10year' allows more detailed visualization of recent events (Fig. 1B). Setting size.scale = 'topology' gives an overview of the relationships among the simulated populations ignoring event times and population sizes (Fig. 1C), which permits demographic events to be displayed in time order even when the time between events is extremely small. There are also options for adjustment of branch positions which can minimize the number of crossings of migration arrows and population branches. One can also plot the population sizes and migrations at a particular point in time (Fig. 1D). The scripts used to run the POPdemog program and generate Figure 1 are provided in the Supplementary Information. The POPdemog program includes an online tutorial which contains examples for different simulators and demographic models, as well as an example showing how migration plots can be overlaid onto a world map.

Funding

Research reported in this publication was supported by the National Human Genome Research Institute of the National Institutes of Health under Award Numbers R01HG005701 and R01HG008359. The content is solely the responsibility of the authors and does not necessarily represent the official views of the National Institutes of Health.

Conflict of Interest: none declared.

References

- Chen,G.K. *et al.* (2009) Fast and flexible simulation of DNA sequence data. *Genome Res.*, **19**, 136–142.
- Hellenthal, G. and Stephens, M. (2007) msHOT: modifying Hudson's ms simulator to incorporate crossover and gene conversion hotspots. *Bioinformatics*, 23, 520–521.
- Hoban, S. et al. (2012) Computer simulations: tools for population and evolutionary genetics. Nat. Rev. Genet., 13, 110–122.
- Hudson, R.R. (2002) Generating samples under a Wright-Fisher neutral model of genetic variation. *Bioinformatics*, 18, 337–338.
- Kelleher, J. et al. (2016) Efficient Coalescent Simulation and Genealogical Analysis for Large Sample Sizes. PLoS Comput. Biol., 12, e1004842.
- Shlyakhter, I. et al. (2014) Cosi2: an efficient simulator of exact and approximate coalescent with selection. Bioinformatics, 30, 3427–3429.
- Staab,P.R. et al. (2015) Scrm: efficiently simulating long sequences using the approximated coalescent with recombination. Bioinformatics, 31, 1680–1682.
- Tennessen, J.A. *et al.* (2012) Evolution and Functional Impact of Rare Coding Variation from Deep Sequencing of Human Exomes. *Science*, 337, 64–69.
- Vernot, B. et al. (2016) Excavating Neandertal and Denisovan DNA from the genomes of Melanesian individuals. Science, 352, 235–239.