

Supporting Information for “[paste title here]”

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Supplemental methods

Generation times needed to explain long-lasting differences between populations

Using the reported generation intervals from WANG *et al.* (2023) (as shown in Figure S4 in their supplemental material), we explored scenarios that could lead to the observed differences between African and non-African populations five to ten thousand generations ago, corresponding to 150–300ka. As discussed in the main text, a 5–10 year difference in generation intervals would require long-lasting structure. Admixture from an unidentified, diverged human lineage has been proposed to explain observed genetic variation in African populations (e.g., HEY *et al.*, 2018; DURVASULA and SANKARARAMAN, 2020; LORENTE-GALDOS *et al.*, 2019, but see RAGSDALE *et al.* (2022)). In such models, a population that was isolated for hundreds of thousands of years contributed 5–10% ancestry to West African populations. (Figure S1). The remaining 90–95% ancestry is shared between present-day Eurasian and West African populations, and this ancestry would have shared historical generation times. While other models of deep population structure in Africa have been proposed, a history of strict isolation between diverged lineages before admixture is more likely to result in a signal of differing ancestral generation times.

In such a scenario, differences in inferred average historical generation times between West African and Eurasian populations must be due to differences in generation times between the two diverged lineages. Using the mutation model WANG *et al.* (2023) inferred from Icelandic pedigree data (JÓNSSON *et al.*, 2017), we modeled the Eurasian mutation spectrum from this time period using paternal and maternal generation times of 20 (18–22, from figure S4 in WANG *et al.*) years. The West African mutation spectrum was modeled as a mixture between this shared spectrum and the mutation spectrum from the diverged lineage, in proportions equal to the admixture proportions. This assumes (1) selection does not strongly influence mutation spectrum proportions, (2) there are no demographic effects such as severe bottlenecks that make mutation spectrum proportions unequal to admixture proportions, and (3) the rates of mutation accumulation along each lineage are similar. Additionally, age- and sex-dependent mutation rates from past populations must match the mutation model from the Icelandic trio data. It is likely that none of these assumptions perfectly hold, but these are the same assumptions in the original inference of generation time histories.

Given the admixture proportion f , inferred West African-ancestral paternal and maternal generation intervals of 28 (27–30) and 23 (22–25) years, and the mutation model $M(p, m)$, we found generation times p_d and m_d in the diverged lineage such that

$$M(28, 23) = (1 - f)M(20, 20) + fM(p_d, m_d).$$

In fitting this model with $f = 0.1$, we found $p_d \approx 92$ and $m_d \approx 48$. If we assume average ancestral paternal and maternal ages in the Eurasian-shared lineage were each 22 years, $p_d \approx 76$ and $m_d \approx 31$. With Eurasian-ancestral intervals of 22 years and $f = 0.2$ (much higher than most inferences), the paternal age would still

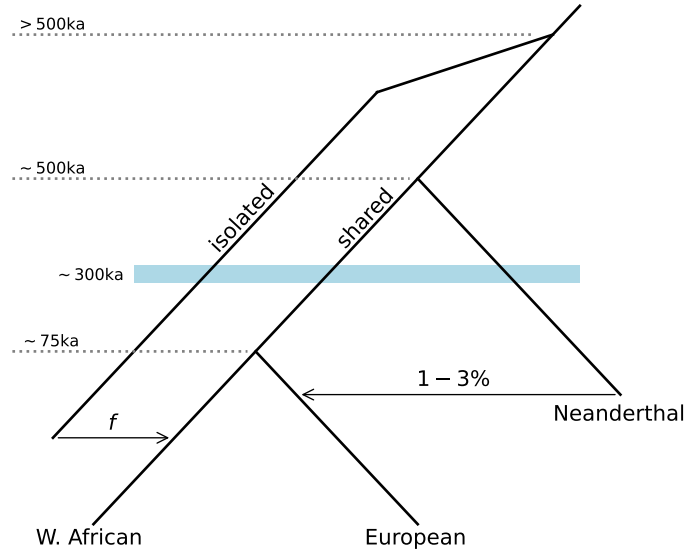


Figure S1: **A model for archaic admixture within Africa.**

need to be over 50 years. From this, we conclude that the generation time history inferred by WANG *et al.* (2023) is incompatible with prevailing models of deep population structure within Africa.

Historical mutation spectra

We followed the filtering choices from WANG *et al.* (2023) in retaining mutations with estimated ages. Namely, triplet mutation contexts associated with a known C→T mutation pulse in Europeans (HARRIS, 2015) and CpG sites were removed. GEVA does not provide allele ages for singletons, but we considered data both with and without singletons from tsdate- and Relate-inferred ages. Variants with allele frequencies greater than 98% were removed to minimize the effect of ancestral-state misidentification.

Variants were binned by age in 100 epochs, divided such that a roughly equal number of variants fell within each bin, as in WANG *et al.* (2023). In most cases, we considered a maximum age of 10,000 generations. Mutation profile trajectories and generation time histories were smoothed using the `loess1d` function from the Python `loess` package, with parameters `frac=0.5` and `degree=2`.

Allele ages from GEVA

Allele ages from Relate

Allele ages from tsdate

References

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- HARRIS, K., 2015 Evidence for recent, population-specific evolution of the human mutation rate. *Proceedings of the National Academy of Sciences* **112**: 3439–3444.
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- WANG, R. J., S. I. AL-SAFFAR, J. ROGERS, and M. W. HAHN, 2023 Human generation times across the past 250,000 years. *Science Advances* **9**: eabm7047.

1 Tables and figures

Dataset	A→C	A→G	A→T	C→A	C→G	C→T
GEVA	0.0946	0.3600	0.0886	0.1201	0.1057	0.2310
tsdate	0.0931	0.3579	0.0899	0.1146	0.1061	0.2384
tsdate (w/singletons)	0.0989	0.3598	0.0908	0.1168	0.1062	0.2275
Relate	0.0991	0.3610	0.0863	0.1124	0.1038	0.2374
Relate (w/singletons)	0.1002	0.3590	0.0921	0.1164	0.1060	0.2263
Trios (phased)	0.0953	0.3649	0.0890	0.0960	0.1216	0.2332
Trios (all mutations)	0.0962	0.3638	0.0923	0.0951	0.1202	0.2324

Table S1: **Mutation profiles from the past 100 generations, compared to Iceland trios.** The most recent time bin for each method included the past ≈ 150 generations. When singletons were included (when using data from **tsdate** and **Relate**), the spectra of estimated recent standing variation were unchanged. Note that **GEVA** does not report ages for singletons. While the three methods provide similar spectra from recent mutations, the spectrum from the Iceland pedigrees differs, in particular for the C→A and C→G classes. These differences are up to 2% of the proportion among all mutations, which corresponds to an under- or over-count of up to $\sim 20\%$ of C→A and C→G mutations, respectively. This difference remains whether the spectrum is estimated from only mutations that were phased in JÓNSSON *et al.* (2017) or from all mutations (phased and unphased).

Dataset	A→C	A→G	A→T	C→A	C→G	C→T
AFR (GEVA)	0.103	0.354	0.094	0.127	0.098	0.224
EAS	0.111	0.341	0.103	0.131	0.094	0.220
EUR	0.102	0.355	0.093	0.125	0.102	0.222
SAS	0.095	0.355	0.090	0.123	0.099	0.238
AFR (Relate)	0.099	0.356	0.084	0.116	0.110	0.236
EAS	0.095	0.359	0.089	0.115	0.097	0.245
EUR	0.100	0.368	0.085	0.110	0.102	0.235
SAS	0.104	0.344	0.090	0.108	0.107	0.246
AFR (tsdate)	0.092	0.354	0.087	0.116	0.110	0.241
EAS	0.098	0.356	0.097	0.112	0.103	0.233
EUR	0.091	0.363	0.089	0.117	0.102	0.238
SAS	0.091	0.359	0.088	0.114	0.107	0.241

Table S2: **Mutation profiles from the past 100 generations in continental population groups.**

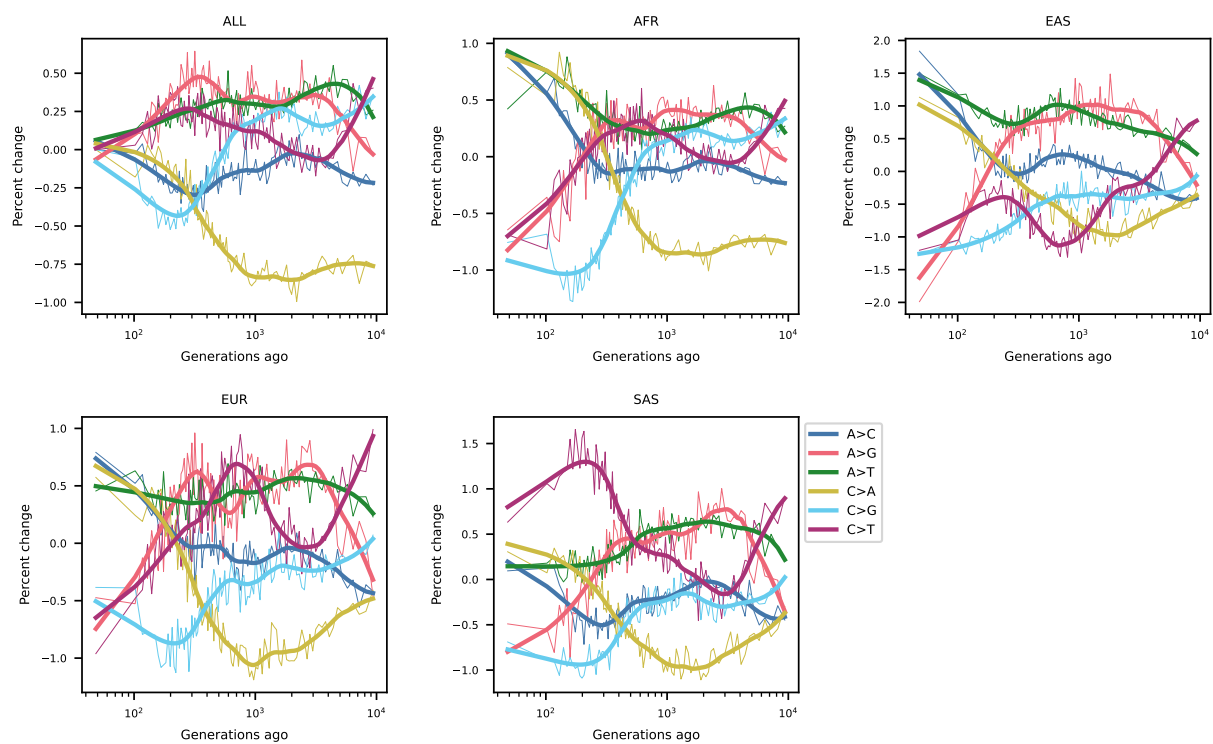


Figure S2: GEVA-inferred mutation spectrum history.

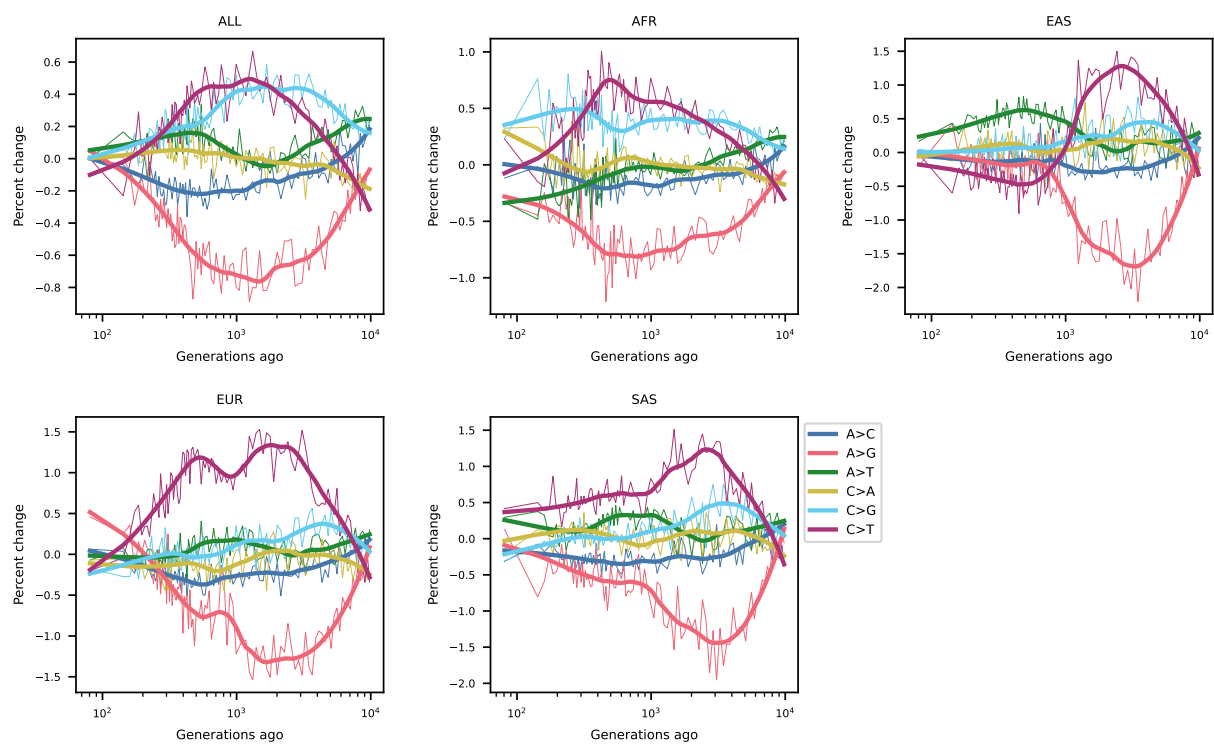


Figure S3: Relate-inferred mutation spectrum history.

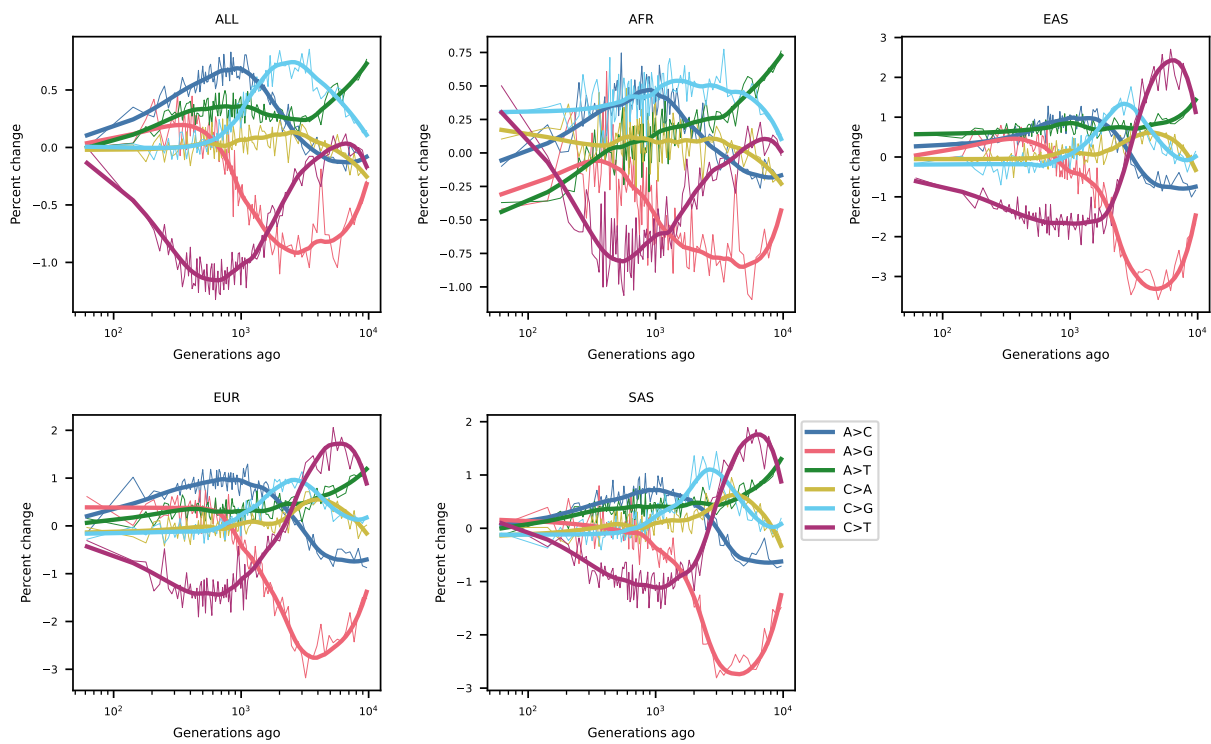


Figure S4: **tsdate-inferred mutation spectrum history.**

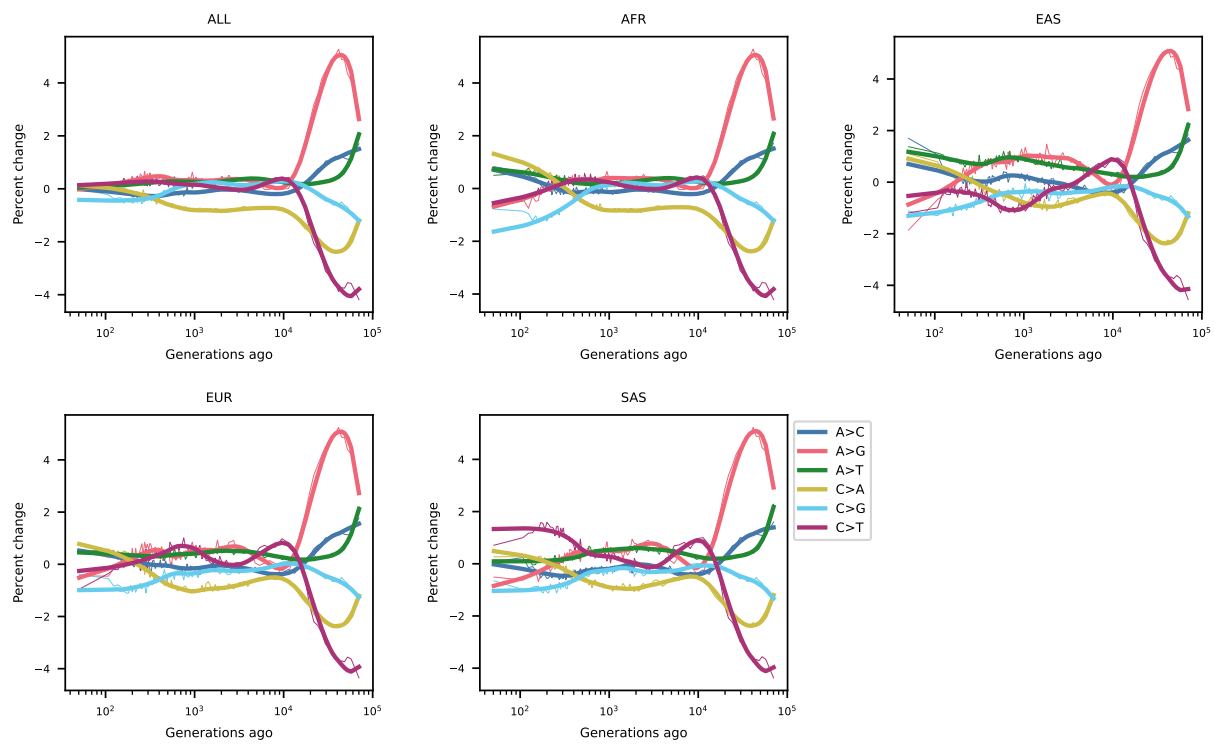


Figure S5: GEVA-inferred mutation spectrum history, extending to 80,000 generations.

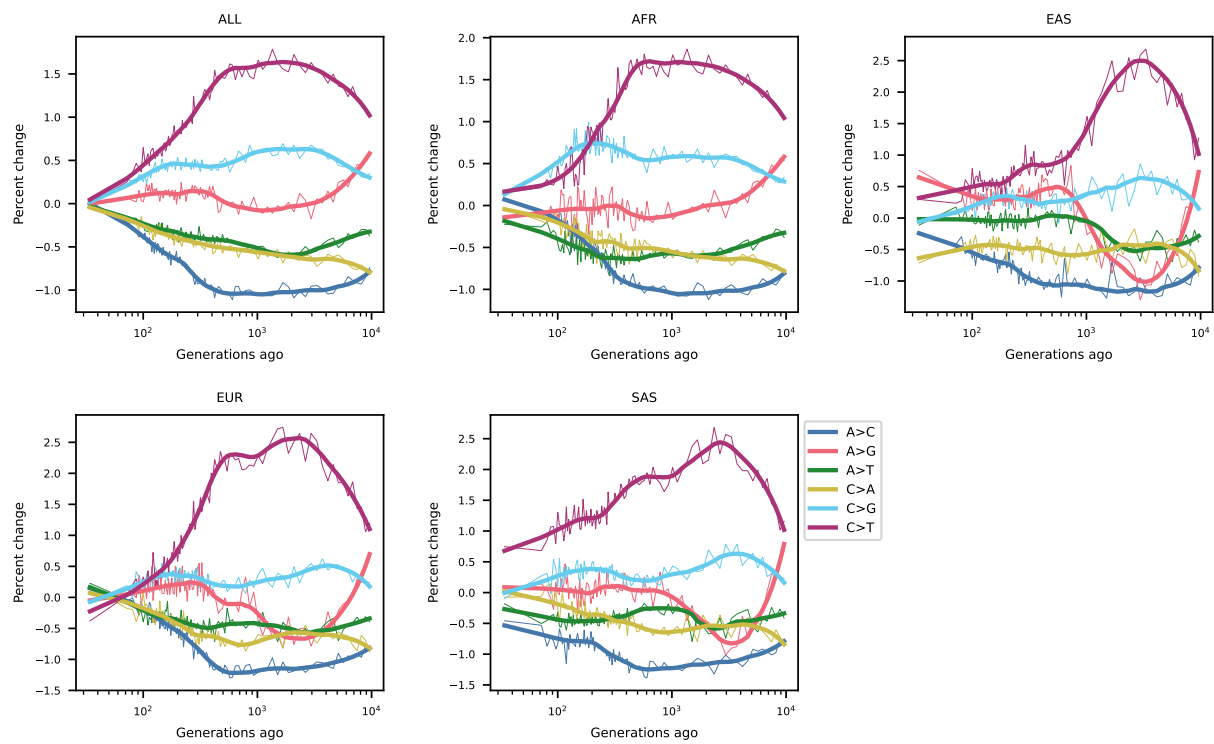


Figure S6: Relate-inferred mutation spectrum history, including singletons.

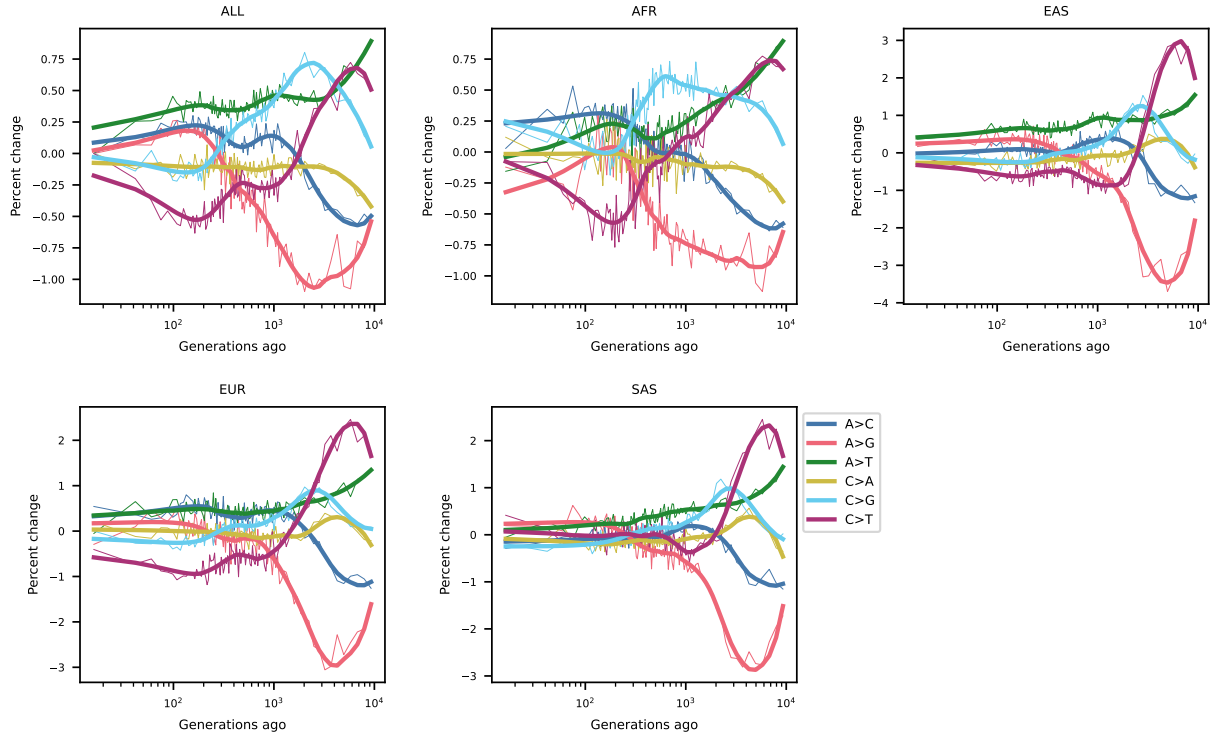


Figure S7: tsdate-inferred mutation spectrum history, including singletons.

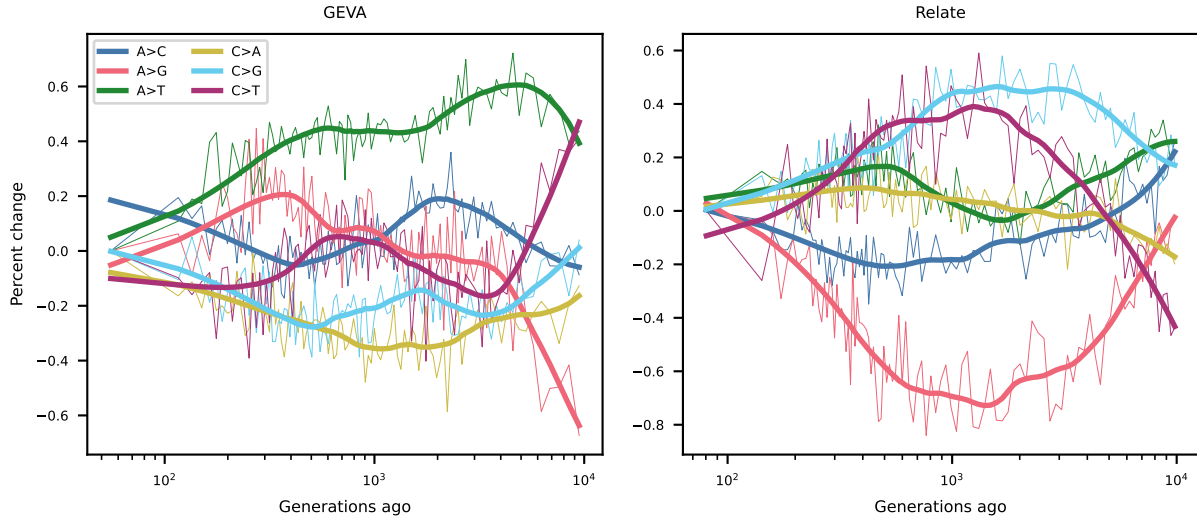


Figure S8: Mutation spectrum histories from mutations that were dated by both GEVA and Relate.

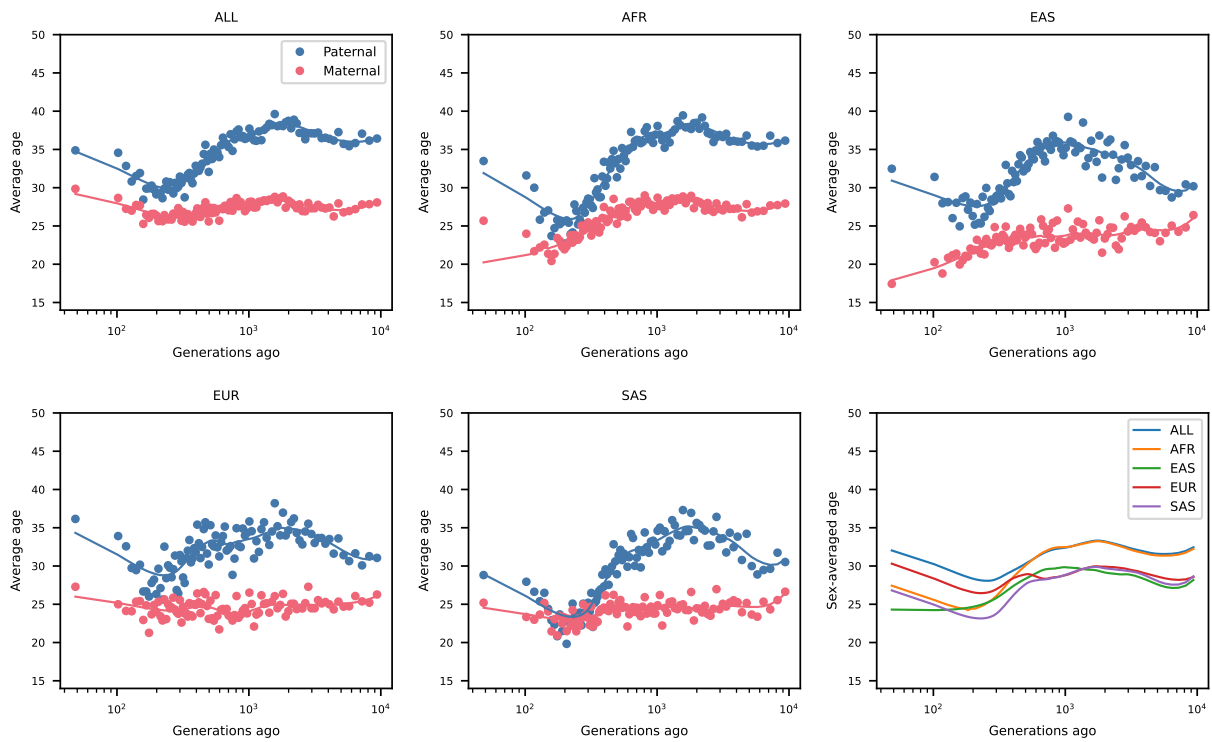


Figure S9: GEVA-inferred generation time histories.

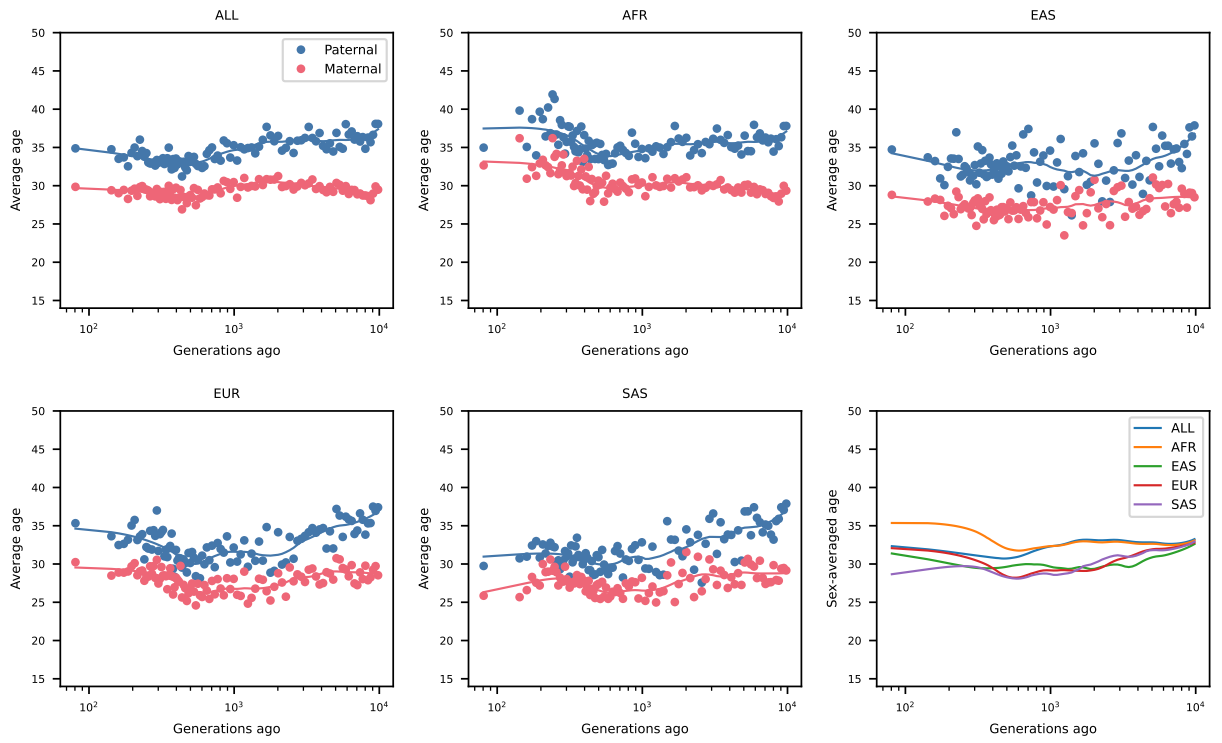


Figure S10: Relate-inferred generation time histories.

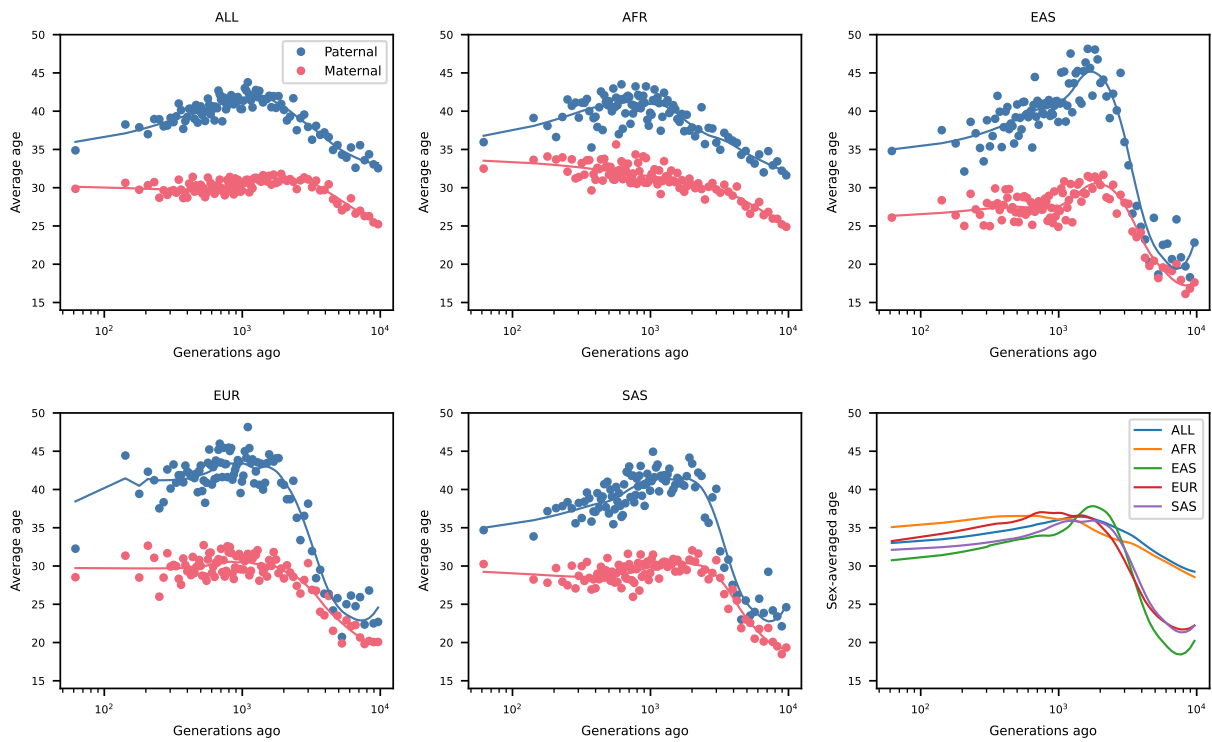


Figure S11: tsdate-inferred generation time histories.

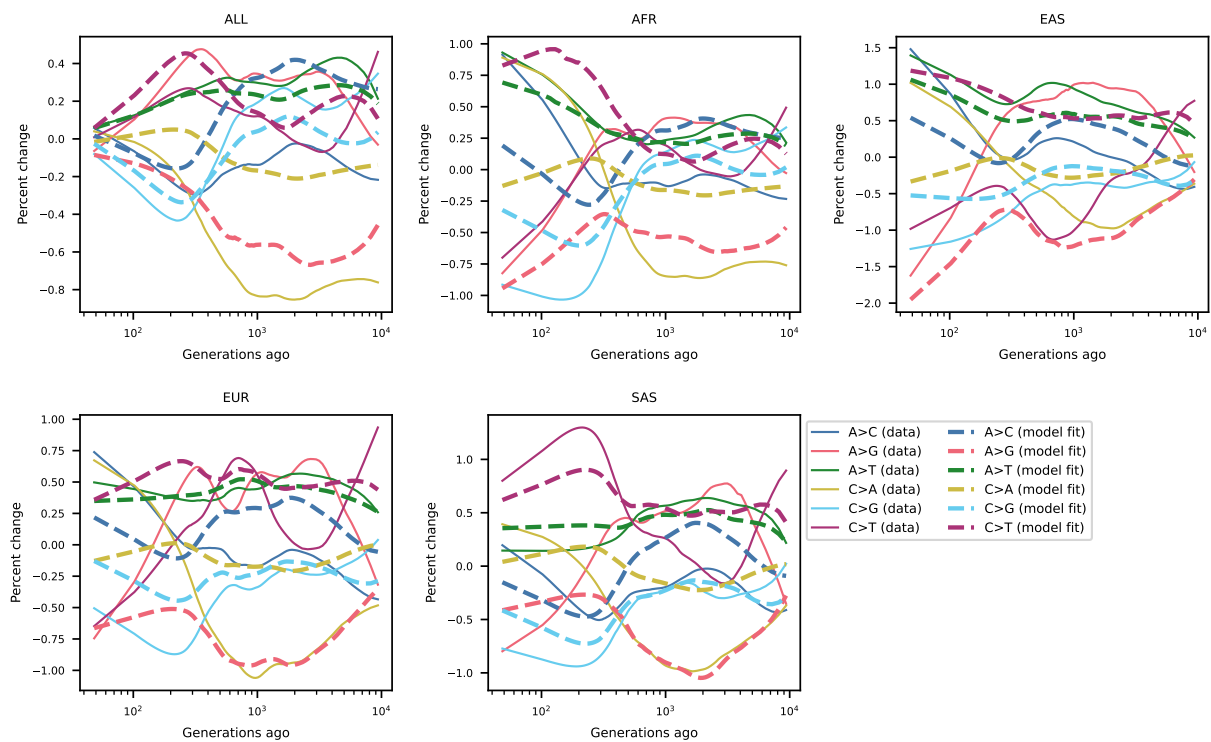


Figure S12: Prediction of mutation spectrum history from GEVA-inferred generation times.

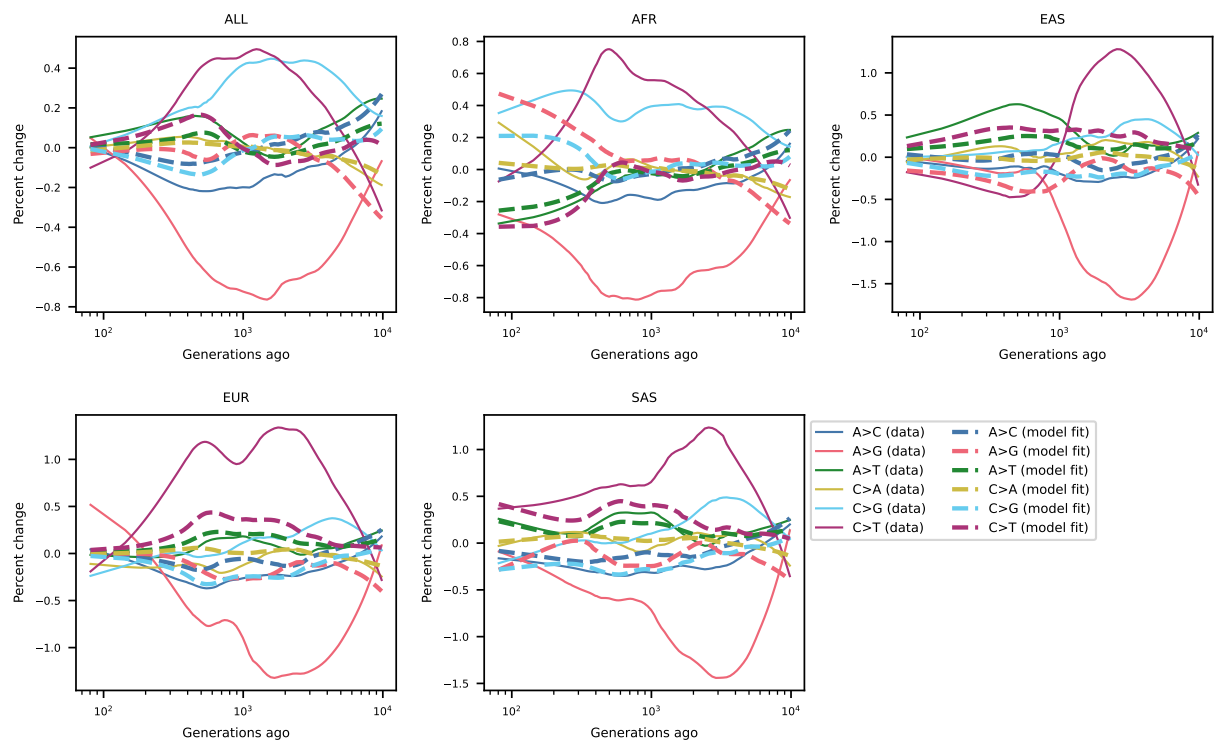


Figure S13: **Prediction of mutation spectrum history from Relate-inferred generation times.**

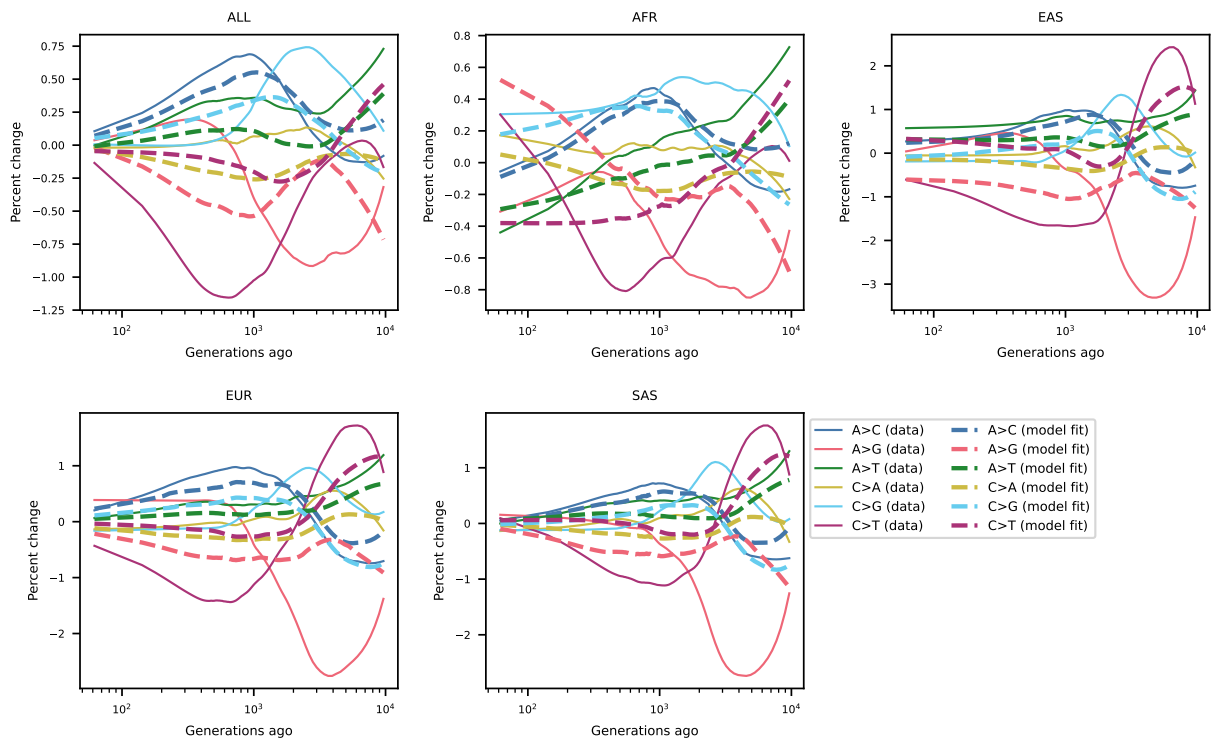


Figure S14: **Prediction of mutation spectrum history from *tsdate*-inferred generation times.**