

# Influence of Dominance and Drift on Lethal Mutations in Human Populations

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This file provides some of the details underlying the simulations used to obtain the figure in the Supplementary Material.

To simulate the Wright-Fisher model for the conventional case of weak selection we use Eq. (6) of the main text, which we reproduce here as

$$X' = \frac{\text{Bin}(2N, X + F(X))}{2N} \quad \begin{array}{l} \text{Wright-Fisher model} \\ \text{for weak selection.} \end{array} \quad (1)$$

For the case of a lethal allele we use Eq. (7) of the main text, which we reproduce here as

$$X' = \frac{\text{Bin}(N, 2X + 2F(X))}{2N} \quad \begin{array}{l} \text{Wright-Fisher model} \\ \text{for a lethal genotype.} \end{array} \quad (2)$$

Here, we just give details for simulation for the conventional case, by directly using Eq. (1) above.

Schematically we proceed as follows.

1. Start with a given value of  $X$ .
2. Using this value of  $X$ , calculate  $X + F(X)$ .
3. Generate a single binomial random number with parameters  $2N$  and  $X + F(X)$  which we write as  $B$   
( $2N$  represents the number of independent trials and  $X + F(X)$  represents the probability of success on each trial).
4. Divide the binomial random number,  $B$ , by  $2N$ , to obtain  $B/(2N)$  which is a relative frequency. This lies in the range 0 to 1.

5. Assign  $X$  to be  $R/(2N)$
6. Store the value of  $X$  and go back to step 1, using the value of  $X$  just obtained, and repeat the procedure.

In this way a set of  $X$ 's is obtained:

$$(X_0, X_1, X_2, \dots).$$

This is a frequency trajectory.

Repeating the entire process again, with the same initial frequency, will generally yield a different trajectory - a replicate trajectory. Statistics of the frequency can be obtained by averaging over replicate trajectories. For example, the mean frequency at generation 2 is estimated by averaging  $X_2$  over a large number of replicate trajectories.

In the file FigureS1.m the implementation of the above in Matlab, for a single replicate trajectory, covering  $T$  generations, is given by

```
for t=1:T
    x=X(t);
    F=((1-h)*u-(h+(2-3*h)*u)*x-(1-2*h)*(1-u)*x^2)/((1+(1-2*h)*u)
        +(1-2*h)*(1-u)*x);
    X(t+1)=binornd(2*N,x+F)/2/N;
end
```