## A mathematical framework for estimating viral transmission fitness and inoculum size using data from a competitive mixtures animal model. Text S1: Results of the simulation studies

Figure 7 shows the recovered estimates for  $\phi$  and N for simulation scenarios D and E described in Table 1 in the main text. Simulations  $D_1$  and  $D_2$  (when compared to both each other and simulation  $A_1$ ) demonstrate that we obtain marginally tighter estimates for  $\phi$  and N by clustering the samples for the donor strain A proportion, p, around either the centre or extreme values. Simulations  $E_1$  and  $E_2$ , where the transmitted inoculum is increased (to 50, up from 4), confirm the expected increase in precision for  $\phi$  due to the lower variability in outcome from the binomial selection process. There is a substantial decrease in precision for the estimate of N.



Figure 7: Recovered estimates for  $\phi$  and N for simulations D and E described in Table 2 in the main text. Note the log scale for the estimate of N. Each boxplot shows the median, 25th and 75th centiles with tails extending to the upper and lower adjacent values and outliers shown as crosses. The asterisk marks the true value used in the simulation. The dashed vertical line is a visual aid to separate simulations D and E. The horizontal line in Figure 7(a) shows a relative transmission fitness of 1. Simulations  $D_1$ ,  $D_2$  and  $E_1$  should be compared to  $A_1$  and  $E_2$  to  $B_4$ .

Figures 8 through 12 present the detailed results for each simulation scenario described in Table 2 in the main text. Each figure shows the numerically determined probability density for the recovered relative transmission fitness  $\phi$  (left) and a histogram for the recovered transmitted inoculum size, N (right). We report the mean and 95% confidence interval for  $\phi$ , and the mode, mean and standard deviation for N.

Evident in Figure 6 in the main text and Figure 7 here, there is a minor bias in the estimate for N. This can be understood by considering  $RSS_M(N)$  in Equation 7 in the main text.  $N = N_{true}$ , where  $N_{true}$  is the true value of the transmitted inoculum size, is the N most likely to generate the minimum difference between  $RSS_D$  and  $RSS_M(N)$  and so the mode of our distribution is accurate. For values of  $N < N_{true}$  the residual must be too large, while for  $N > N_{true}$  the residual, while expected to be too small, may be too large. It follows that a slight right bias will be found in the histogram for N.



(a) Simulation  $A_1: \phi_{\rm est} = 1.14,$  (0.47, 2.40), mode( $N_{\rm est}) = 4,$  mean( $N_{\rm est}) = 7.3,$   $\sigma(N_{\rm est}) = 5.5$ 



(b) Simulation  $A_2$ :  $\phi_{\rm est} = 0.98$ , (0.38, 2.02), mode $(N_{\rm est}) = 4$ , mean $(N_{\rm est}) = 7.5$ ,  $\sigma(N_{\rm est}) = 6.2$ 



(c) Simulation A<sub>3</sub>:  $\phi_{\text{est}} = 0.87$ , (0.36, 1.83), mode $(N_{\text{est}}) = 4$ , mean $(N_{\text{est}}) = 7.3$ ,  $\sigma(N_{\text{est}}) = 5.5$ 



(d) Simulation A<sub>4</sub>:  $\phi_{\text{est}} = 0.55$ , (0.22, 1.09), mode( $N_{\text{est}}$ ) = 4, mean( $N_{\text{est}}$ ) =  $\sigma(N_{\text{est}}) = 5.7$ 

Figure 8: Simulation A: Fits to  $\phi$  (left) and N (right)



(a) Simulation  $B_1: \phi_{\rm est}=1.07,$  (0.70, 1.57),  ${\rm mode}(N_{\rm est})=4,$   ${\rm mean}(N_{\rm est})=4.9,$   $\sigma(N_{\rm est})=1.9$ 



(b) Simulation  $B_2$ :  $\phi_{\rm est}=0.91,$  (0.58, 1.37), mode( $N_{\rm est})=4,$  mean( $N_{\rm est})=4.9,$   $\sigma(N_{\rm est})=1.9$ 



(c) Simulation  $B_3$ :  $\phi_{\rm est}=0.81,$  (0.52, 1.22), mode( $N_{\rm est})=4,$  mean( $N_{\rm est})=4.9,$   $\sigma(N_{\rm est})=2.0$ 



(d) Simulation  $B_4$ :  $\phi_{\rm est}=0.51,$  (0.32, 0.78), mode( $N_{\rm est})=4,$  mean( $N_{\rm est})=4.9,$   $\sigma(N_{\rm est})=2.0$ 

Figure 9: Simulation B: Fits to  $\phi$  (left) and N (right)



(a) Simulation  $C_1: \ \phi_{\rm est} = 1.06,$  (0.83, 1.34), mode( $N_{\rm est}) = 4,$  mean( $N_{\rm est}) = 5.0,$   $\sigma(N_{\rm est}) = 0.94$ 



(b) Simulation  $C_2$ :  $\phi_{\rm est}=0.91,$  (0.70, 1.15),  ${\rm mode}(N_{\rm est})=4,$   ${\rm mean}(N_{\rm est})=4.3,$   $\sigma(N_{\rm est})=0.96$ 



(c) Simulation  $C_3$ :  $\phi_{\rm est}=0.81,$  (0.62, 1.03), mode( $N_{\rm est})=4,$  mean( $N_{\rm est})=4.3,$   $\sigma(N_{\rm est})=0.96$ 



Figure 10: Simulation C: Fits to  $\phi$  (left) and N (right)



(a) Simulation  $D_1$ :  $\phi_{\text{est}} = 1.07$ , (0.74, 1.47), mode $(N_{\text{est}}) = 4$ , mean $(N_{\text{est}}) = 4.8$ ,  $\sigma(N_{\text{est}}) = 1.7$ 



(b) Simulation  $D_2$ :  $\phi_{\text{est}} = 1.09$ , (0.62, 1.85), mode $(N_{\text{est}}) = 4$ , mean $(N_{\text{est}}) = 5.0$ ,  $\sigma(N_{\text{est}}) = 2.1$ 

Figure 11: Simulation D: Fits to  $\phi$  (left) and N (right)



(a) Simulation  $E_1$ :  $\phi_{\text{est}} = 1.06$ , (0.83, 1.33), mode $(N_{\text{est}}) = 46$ , mean $(N_{\text{est}}) = 94.4$ ,  $\sigma(N_{\text{est}}) = 79.4$ 



(b) Simulation  $E_2$ :  $\phi_{\text{est}} = 0.50$ , (0.44, 0.57), mode $(N_{\text{est}}) = 48$ , mean $(N_{\text{est}}) = 60.8$ ,  $\sigma(N_{\text{est}}) = 25.3$ 

Figure 12: Simulation E: Fits to  $\phi$  (left) and N (right)