## S1 Model: Non-neutral models

## **Deleterious model**

Let  $s_d$  be the selection coefficient of the deleterious mutation (where the fitness of a cell that is homoplasmic for mutant mitochondria is  $1 - s_d$ ). Assuming that the cost imposed by mutant mitochondria can be modeled by a concave function, the fitness of a cell with *i* mutant mitochondria is given by

$$w_d(i) = w(i) \left( 1 - s_d \left( \frac{i}{n} \right)^2 \right)$$
, where  $w(i)$  is determined by either equation (1) or (3).

## Advantageous model

Again, we generate w(i) using equation (1) or equation (3). We let  $s_a$  be the selection coefficient of the deleterious mutation (where the fitness of a cell that is homoplasmic for wild type mitochondria is  $1 - s_a$ ). Since there are no data as to how fitness increases as advantageous mitochondria accumulate, we model the increase in fitness using a concave function and a convex function. If we assume that advantageous mutations convey fitness benefits consistent with a concave function, the fitness of a cell with *i* mutant mitochondria is given by

$$w_a(i) = w(i) \left( 1 - s_a \left( \frac{n-i}{n} \right)^2 \right).$$

If we assume that fitness accumulates as a convex function of the number of mutant mitochondria, then the fitness of a cell with *i* mutant mitochondria is given by

$$w_a(i) = w(i) \left( 1 - s_d \sqrt{\frac{n-i}{n}} \right).$$