

## Supplementary Table 2

Overview of kinetic constants used for the construction of the model.

Enzyme	EC number	Kinetic parameter	References	Rate Law
NADA	3.5.1.19	$K_M:9.6\mu\text{M}$ $K_{iP}:120\mu\text{M}$ $k_{cat}:0.65s^{-1}$	[1]	Product inhibition
NADS	6.3.5.1	$K_M:190\mu\text{M}$ $k_{cat}:21s^{-1}$	[2]	HMM
NMNAT	2.7.7.1 2.7.7.18	$K_{M_{NaMN}}:67.7\mu\text{M}$ $k_{cat_{NaMN}}:42.9s^{-1}$ $K_{M_{NMN}}:22.3\mu\text{M}$ $k_{cat_{NMN}}:53.8s^{-1}$ $K_{M_{NMN}}:59\mu\text{M}$ $k_{cat_{NAD}}:129.1s^{-1}$ $K_{M_{NaAD}}:502\mu\text{M}$ $k_{cat_{NaAD}}:103.8s^{-1}$	[3] <sup>1</sup>     [4] <sup>2</sup>  [4] <sup>3</sup>	Substrate Competition
NMNT	2.1.1.1	$K_M:400\mu\text{M}$ $K_{iP}:60\mu\text{M}$ $k_{cat}:8.1s^{-1}$	[5]  [6]	Product inhibition
NamPT	6.3.5.1	$K_M:5\text{nM}$ $k_{cat}:0.0077s^{-1}$ $K_{i_{NAD}}:2.1\mu\text{M}$	[7]	Competitive inhibition
NAPRT	2.4.2.11	$K_M:1.5\mu\text{M}$ $k_{cat}:3.3s^{-1}$	[7]	HMM
SIRT1	3.5.1.-	$K_M:29\mu\text{M}$ $K_{iP}:60\mu\text{M}$ $k_{cat}:0.67s^{-1}$	[8]	Product inhibition
NT5	3.1.3.5	$K_{M_{NaMN}}:3.5\text{mM}$ $k_{cat_{NaMN}}:2.8s^{-1}$ $K_{M_{NMN}}:5\text{mM}$ $k_{cat_{NMN}}:0.5s^{-1}$	[9]	HMM
PNP	2.4.2.1	$K_M:1.48\text{mM}$ $k_{cat}:40s^{-1}$	[10]	HMM
NRK	2.7.1.173	$K_M:3.4\mu\text{M}$ $k_{cat}:0.23s^{-1}$	[11]	HMM

<sup>1</sup>Values for NMNAT1 used

<sup>2</sup>Keq used for calculation of turnover rate of reverse reaction

<sup>3</sup>Equilibrium constant used for calculation of turnover rate of reverse reaction

## Amount of enzymes and import rates

The total enzyme concentration was set to 10 for all enzymes except NamPT and NMNAT, for which the concentration was set to 100. Concentration of potential co-substrate was assumed to be constant and not-limiting for the reaction. Thus being implicitly represented by turnover rates measured at saturating conditions for the cosubstrate. Nam import rates for import into the system was set to 0.1  $\mu\text{M}/\text{ls}$  for all simulations. For the two compartment simulation compartment size was equal to ... for both compartments. The Nam import rates were set to 100/s for both compartments. The amount of NADA present was set to 100. Thus equal to the amount of NamPT used.

## Kinetic Rate Laws

### Product Inhibition

$$v = \frac{E_T \cdot k_{cat} \cdot S}{K_M + S + \frac{K_M \cdot P}{K_{iP}}} \quad (1)$$

### Competitive Inhibition

$$v = \frac{E_T \cdot k_{cat} \cdot S}{K_M + S + \frac{K_M \cdot I}{K_{iI}}} \quad (2)$$

### Henry-Michaelis Menten for irreversible reactions (HMM)

$$v = \frac{E_T \cdot k_{cat} \cdot S}{K_M + S} \quad (3)$$

### Substrate Competition at NMNAT

$$v = E_T \cdot \frac{\frac{k_{cat_A} \cdot A \cdot B}{K_{MA}} - \frac{k_{cat_P} \cdot P \cdot Q}{K_{MP}}}{1 + \frac{A}{K_{MA}} + \frac{B}{K_{MB}} + \frac{P}{K_{MP}} + \frac{Q}{K_{MQ}}} \quad (4)$$

## References

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