

Supplementary Table 1

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] ¹ [4] ² [4] ³	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
NamPRT	6.3.5.1	$K_M:5\text{nM}$ $k_{cat}:0.0077s^{-1}$	[7]	HMM
NAPRT	2.4.2.11	$K_M:23\mu\text{M}$ $k_{cat}:3.3s^{-1}$	[8]	HMM
SIRT1	3.5.1.-	$K_M:29\mu\text{M}$ $K_{iP}:60\mu\text{M}$ $k_{cat}:0.67s^{-1}$	[9]	Product inhibition
NT5 ⁴	3.1.3.5	$K_M:100\mu\text{M}$ $k_{cat}:10s^{-1}$ ⁶	[10] ⁵	HMM

The total enzyme concentration was set to 10 for all enzymes if not mentioned otherwise. Concentration of potential cosubstrate was assumed to be constant and considered to be not limiting the reaction and thus represented by the maximal velocities given.

¹Values for NMNAT1 used

²Keq used for calculation of turnover rate of reverse reaction

³Equilibrium constant used for calculation of turnover rate of reverse reaction

⁴As parameter values for NRK and PNP were not available we omitted these enzymes and simulated NA and NAM as direct products

⁵approx. IC50 value for NAD used

⁶Average value for pyrimidines used

Kinetic Rate Laws

Product Inhibition

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

Henry-Michaelis Menten for irreversible reactions (HMM)

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

Substrate Competition at NMNAT

$$v = E_T \cdot \frac{\frac{k_{cat_A} \cdot A \cdot B}{K_{M_A}} - \frac{k_{cat_P} \cdot P \cdot Q}{K_{M_P}}}{1 + \frac{A}{K_{M_A}} + \frac{B}{K_{M_B}} + \frac{P}{K_{M_P}} + \frac{Q}{K_{M_Q}}} \quad (3)$$

References

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