Additional Table 2 - Kinetic constants of the enzymes in *M. tuberculosis* model-1 and model-2. Vmax refers to maximal rate of an enzymatic reaction.  $K_{S1}$  and  $K_{S2}$  refer to the  $K_M$  values of substrates, and,  $K_{P1}$  and  $K_{P2}$  refer to the  $K_M$  values of products of the reactions. They are numbered in the order they appear in the chemical equations shown in the table.

Reaction	Enzyme	Abbreviation	Vmax (in nmol/min/mg protein)	Vmax (mM/min)	K <sub>S1</sub> (mM)	K <sub>S2</sub> (mM)	K <sub>P1</sub> (mM)	$K_{P2}$ (mM)
ACA + OAA =	citrate synthase	CS	405 [1]	64.8	0.05 [2]	$0.012^{a}[2]$	0.5	0.12
COA + CIT								
CIT = ICIT	aconitase	ACN	195 [1]	31.2	1.7 <sup>b</sup> [2]		$0.7^{c}[2]$	
ICIT = aKG	<i>iso</i> citrate dehydrogenase 1	ICD1	64 [1]	10.2	0.03 [2]		0.3	
ICIT = aKG	<i>iso</i> citrate dehydrogenase 2	ICD2	-	9.965 <sup>d</sup>	0.06 <sup>d</sup>		0.6	
$aKG = SCA^{e}$	α-ketoglutarate dehydrogenase complex	KDH	-	57.344 <sup>f</sup>	0.1 <sup>f</sup>		1	
SCA = SUC	succinyl-CoA synthetase	ScAS	7.5 [1]	1.2	0.02 <sup>g</sup> [2]		5 <sup>h</sup> [2]	
aKG = SSA	α-ketoglutarate decarboxylase	KGD	-	48.3 <sup>i</sup>	0.48 [1]		4.8	
SSA = SUC	succinic semialdehyde dehydrogenase	SSADH	40.7 [1]	6.51	0.015 <sup>j</sup> [2]		0.15	
SUC = FA	succinate dehydrogenase	SDH	6.4 [1]	1.02	0.15 <sup>k</sup> [2]		0.12 <sup>k</sup> [2]	
FA = MAL	fumarase	FUM	548 [1]	87.7	$0.25^{1}[2]$		$2.38^{1}[2]$	
MAL = OAA	malate dehvdrogenase	MDH	1149 [1]	184	0.833 <sup>k</sup> [2]		0.0443 <sup>k</sup> [2]	
ICIT = SUC + GLY	isocitrate lyase 1	ICL1	1.83 [3]	1.172 <sup>m</sup>	0.145 [2]		0.59 <sup>f</sup>	0.13 <sup>f</sup>
ICIT = SUC + GLY	isocitrate lyase 2	ICL2	-	0.391 <sup>n</sup>	1.3 [4]		5.9°	1.3°
GLY + ACA = MAL + COA	malate synthase	MS	125 [3]	20	0.057 [2]	0.03 [2]	$1^{\mathrm{f}}$	$0.1^{\mathrm{f}}$

## Notes for additional table 2

<sup>a</sup>Value taken from *Bacillus megaterium* 

<sup>b</sup>Value taken from *Bacillus subtilis* 

<sup>c</sup>Value taken from *Bacillus cereus* 

1

Additional file 2 to the paper titled "Kinetic modeling of tricarboxylic acid cycle and glyoxylate bypass in *Mycobacterium tuberculosis*, and its application to assessing drug targets" submitted by Vivek Kumar Singh and Indira Ghosh

<sup>d</sup>Banerjee et. al [5] estimated the Vmax of purified ICD1 and ICD2, and Tian et. al [1] estimated the Vmax of ICD1 in cell lysate. Using the ratio of Vmax of purified ICD2 to Vmax of purified ICD1, and the Vmax of ICD1 in cell lysate, the Vmax of ICD2 in cell lysate was estimated. Similar logic was used to estimate the K<sub>M</sub> value of ICIT for ICD2

eThis reaction is present only in M. tuberculosis model-1

<sup>f</sup>Value taken from *E. coli* 

<sup>g</sup>Value taken from *Calliphoridae* 

<sup>h</sup>Value taken from *Nitrosomonas europaea* 

<sup>i</sup>Value was not available in cell lysate condition, so average of the Vmax of all TCA cycle enzymes was used

<sup>j</sup>Value taken from *Sus scrofa* 

<sup>k</sup>Value taken from *Mycobacterium phlei* 

<sup>1</sup>Value taken from bacterium

<sup>m</sup>Multiply the calculated Vmax (=0.293 mM/min) by 4 because ICL activity increases 4-fold in persistent mycobacteria [3]

<sup>n</sup>Logic similar to that used for the estimation of Vmax for ICD2 was used to estimate Vmax for ICL2 and K<sub>M</sub> value of ICIT for ICL2. The data on purified ICL1 and ICL2 were obtained from Bentrup et. al [4])

<sup>o</sup>Since the K<sub>M</sub> of ICIT for ICL2 is 10 times the K<sub>M</sub> of ICIT for ICL1 [4], the K<sub>M</sub> of the products i.e. SUC and GLY were also increased 10 fold.

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## Abbreviations of metabolites

- ACA acetyl-CoA
- OAA oxaloacetate
- COA CoA
- CIT citrate
- ICIT isocitrate
- aKG α-ketoglutarate
- SCA succinyl-CoA
- SUC succinate
- FA fumarate
- MAL malate
- GLY glyoxylate
- SSA succinic semialdehyde

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