

Table S1 Metabolic reaction list for *E. coli*

No	Gene	Reaction
1	Biomass formation	0.14176 Glyc3P + 26.2949 ATP + 0.60097 Ala + 0.10124 Cys + 0.26647 Asp + 0.30747 Glu + 0.2048 Phe + 0.67725 Gly + 0.10473 His + 0.32116 Ile + 0.37935 Lys + 0.49804 Leu + 0.16989 Met + 0.26647 Asn + 0.24436 Pro + 0.29091 Gln + 0.32698 Arg + 0.38031 Ser + 0.28044 Thr + 0.46778 Val + 0.062835 Trp + 0.15244 Tyr + 0.1489 rATP + 0.18319 rGTP + 0.11366 rCTP + 0.12273 rUTP + 0.023904 dATP + 0.024582 dGTP + 0.024582 dCTP + 0.023904 dTTP + 0.28352 avg_FS + 0.0069264 UDPGlc + 0.010368 CDPEth + 0.010368 OH_myr_ac + 0.010368 C14_0_FS + 0.010368 CMP_KDO + 0.010368 NDPHep + 0.0069264 TDPGlc + 0.01656 UDP_NAG + 0.01656 UDP_NAM + 0.01656 di_am_pim + 0.0924 ADPGlc ==> Biomass
2	Nitrogen uptake	==> N
3	CO2 exchange	CO2 <=>
4	Sulfur uptake	4 ATP + 4 NADPH ==> S
5	<i>pts</i>	PEP + GLC ==> G6P + Pyr
6	<i>glk</i>	ATP + GLC ==> G6P
7	Succinate exchange	Succ ==>
8	<i>gps</i>	DHAP + NADH <=> Glyc3P
9	Lactate exchange	Lac ==>
10	Ethanol exchange	Eth ==>
11	Acetate exchange	Ac ==>
12	Formate exchange	Form ==>
13	<i>pgi</i>	G6P <=> F6P
14	<i>fbp</i>	F16P ==> F6P
15	<i>fba</i>	F16P <=> DHAP + G3P
16	<i>tpi</i>	DHAP <=> G3P
17	<i>gap</i>	G3P <=> DPG + NADH
18	<i>pgk</i>	DPG <=> 3PG + ATP
19	<i>Gpm</i>	3PG <=> 2PG
20	<i>eno</i>	2PG <=> PEP
21	<i>pyk</i>	PEP ==> Pyr + ATP
22	<i>pps</i>	Pyr + 2 ATP ==> PEP
23	<i>lpd</i>	Pyr ==> AcCoA + NADH + CO2
24	<i>glt</i>	AcCoA + OxA ==> Cit
25	<i>acn</i>	Cit <=> ICit
26	<i>icd</i>	ICit <=> alKG + NADPH + CO2
27	<i>sucAB</i>	alKG ==> SuccCoA + NADH + CO2
28	<i>sucCD</i>	SuccCoA <=> Succ + ATP
29	<i>sdh</i>	Succ ==> Fum + QuiH2
30	<i>frd</i>	Fum + QuiH2 ==> Succ
31	<i>Fum</i>	Fum <=> Mal
32	<i>mdh</i>	Mal <=> OxA + NADH
33	<i>aceA</i>	ICit ==> Succ + Glyox
34	<i>aceB</i>	AcCoA + Glyox ==> Mal
35	<i>zwf</i>	G6P <=> PGlac + NADPH
36	<i>adh</i>	AcCoA + NADH <=> Adh

Table S1 (continued)

37	<i>adh</i>	$\text{NADH} + \text{Adh} \rightleftharpoons \text{Eth}$
38	<i>pgl</i>	$\text{PGlac} \implies \text{PGLuc}$
39	<i>gnd</i>	$\text{PGLuc} \implies \text{R15P} + \text{NADPH} + \text{CO}_2$
40	<i>rpe</i>	$\text{R15P} \rightleftharpoons \text{X5P}$
41	<i>rpi</i>	$\text{R15P} \rightleftharpoons \text{R5P}$
42	<i>tktAB</i>	$\text{R5P} + \text{X5P} \rightleftharpoons \text{G3P} + \text{S7P}$
43	<i>tal</i>	$\text{G3P} + \text{S7P} \rightleftharpoons \text{F6P} + \text{E4P}$
44	<i>tktAB</i>	$\text{E4P} + \text{X5P} \rightleftharpoons \text{F6P} + \text{G3P}$
45	<i>edd</i>	$\text{PGLuc} \implies \text{KetoPGLuc}$
46	<i>eda</i>	$\text{KetoPGLuc} \rightleftharpoons \text{G3P} + \text{Pyr}$
47	<i>pck</i>	$\text{OxA} + \text{ATP} \implies \text{PEP} + \text{CO}_2$
48	<i>ppc</i>	$\text{PEP} + \text{CO}_2 \implies \text{OxA}$
49	<i>pta</i>	$\text{AcCoA} \rightleftharpoons \text{AcP}$
50	<i>ack</i>	$\text{AcP} \rightleftharpoons \text{ATP} + \text{Ac}$
51	<i>pfl</i>	$\text{Pyr} \implies \text{AcCoA} + \text{Form}$
52	<i>ldh</i>	$\text{Pyr} + \text{NADH} \rightleftharpoons \text{Lac}$
53	<i>nuo</i>	$\text{NADH} \rightleftharpoons \text{QuiH}_2 + 2 \text{H}_{\text{ex}}$
54	<i>pntA</i>	$\text{NADH} + \text{H}_{\text{ex}} \rightleftharpoons \text{NADPH}$
55	ATP Synthesis	$3 \text{H}_{\text{ex}} \rightleftharpoons \text{ATP}$
56	ATPdrain	$\text{ATP} \implies$
57	<i>aro</i>	$2 \text{PEP} + \text{E4P} + \text{ATP} + \text{NADPH} \implies \text{Chor}$
58	<i>prsA</i>	$\text{R5P} + 2 \text{ATP} \implies \text{PRPP}$
59	<i>met</i>	$\text{ATP} + \text{NADPH} \rightleftharpoons \text{MTHF}$
60	<i>alaB</i>	$\text{Pyr} + \text{Glu} \implies \text{alKG} + \text{Ala}$
61	<i>avt</i>	$2 \text{Pyr} + \text{NADPH} + \text{Glu} \implies \text{alKG} + \text{CO}_2 + \text{Val}$
62	<i>ilv</i>	$2 \text{Pyr} + \text{AcCoA} + \text{NADPH} + \text{Glu} \implies \text{alKG} + \text{NADH} + 2 \text{CO}_2 + \text{Leu}$
63	<i>asn</i>	$2 \text{ATP} + \text{N} + \text{Asp} \implies \text{Asn}$
64	<i>asp</i>	$\text{OxA} + \text{Glu} \implies \text{alKG} + \text{Asp}$
65	<i>Lys</i>	$\text{di}_{\text{am}}_{\text{pim}} \implies \text{CO}_2 + \text{Lys}$
66	<i>met</i>	$\text{SuccCoA} + \text{ATP} + 2 \text{NADPH} + \text{MTHF} + \text{Cys} + \text{Asp} \implies \text{Pyr} + \text{Succ} + \text{N} + \text{Met}$
67	<i>thr</i>	$2 \text{ATP} + 2 \text{NADPH} + \text{Asp} \implies \text{Thr}$
68	<i>ilv</i>	$\text{Pyr} + \text{NADPH} + \text{Glu} + \text{Thr} \implies \text{alKG} + \text{CO}_2 + \text{N} + \text{Ile}$
69	<i>his</i>	$\text{ATP} + \text{PRPP} + \text{Gln} \implies \text{alKG} + 2 \text{NADH} + \text{His}$
70	<i>gab</i>	$\text{alKG} + \text{NADPH} + \text{N} \implies \text{Glu}$
71	<i>gln</i>	$\text{ATP} + \text{N} + \text{Glu} \implies \text{Gln}$
72	<i>pro</i>	$\text{ATP} + 2 \text{NADPH} + \text{Glu} \implies \text{Pro}$
73	<i>arg</i>	$\text{AcCoA} + 4 \text{ATP} + \text{NADPH} + \text{CO}_2 + \text{N} + \text{Asp} + 2 \text{Glu} \implies \text{alKG} + \text{Fum} + \text{Ac} + \text{Arg}$
74	<i>trp</i>	$\text{Chor} + \text{PRPP} + \text{Gln} + \text{Ser} \implies \text{G3P} + \text{Pyr} + \text{CO}_2 + \text{Glu} + \text{Trp}$
75	<i>tyr</i>	$\text{Chor} + \text{Glu} \implies \text{alKG} + \text{NADH} + \text{CO}_2 + \text{Tyr}$
76	<i>phe, tyr</i>	$\text{Chor} + \text{Glu} \implies \text{alKG} + \text{CO}_2 + \text{Phe}$
77	<i>ser</i>	$3 \text{PG} + \text{Glu} \implies \text{alKG} + \text{NADH} + \text{Ser}$
78	<i>gly</i>	$\text{Ser} \implies \text{MTHF} + \text{Gly}$
79	<i>cys</i>	$\text{AcCoA} + \text{S} + \text{Ser} \implies \text{Ac} + \text{Cys}$
80	rATP_Synth	$5 \text{ATP} + \text{CO}_2 + \text{PRPP} + 2 \text{MTHF} + 2 \text{Asp} + \text{Gly} + 2 \text{Gln} \implies 2 \text{Fum} + \text{NADPH} + 2 \text{Glu} + \text{rATP}$

Table S1 (continued)

81	rGTP_Synth	6 ATP + CO ₂ + PRPP + 2 MTHF + Asp + Gly + 3 Gln ==> 2 Fum + NADH + NADPH + 3 Glu + rGTP
82	rCTP_Synth	ATP + Gln + rUTP ==> Glu + rCTP
83	rUTP_Synth	4 ATP + N + PRPP + Asp ==> NADH + rUTP
84	dATP_Synth	NADPH + rATP ==> dATP
85	dGTP_Synth	NADPH + rGTP ==> dGTP
86	dCTP_Synth	NADPH + rCTP ==> dCTP
87	dTTP_Synth	2 NADPH + MTHF + rUTP ==> dTTP
88	avg_FS_Synth	8.24 AcCoA + 7.24 ATP + 13.91 NADPH ==> avg_FS
89	UDPGlc_Synth	G6P + ATP ==> UDPGlc
90	CDPEth_Synth	3PG + 3 ATP + NADPH + N ==> NADH + CDPEth
91	OH_myrc_ac_Synth	7 AcCoA + 6 ATP + 11 NADPH ==> OH_myrc_ac
92	C14_0_FS_Synth	7 AcCoA + 6 ATP + 12 NADPH ==> C14_0_FS
93	CMP_KDO_Synth	PEP + R5P + 2 ATP ==> CMP_KDO
94	NDPHep_Synth	1.5 G6P + ATP ==> 4 NADPH + NDPHep
95	TDPGlcS_Synth	F6P + 2 ATP + N ==> TDPGlcS
96	UDP_NAG_Synth	F6P + AcCoA + ATP + Gln ==> Glu + UDP_NAG
97	UDP_NAM_Synth	PEP + NADPH + UDP_NAG ==> UDP_NAM
98	di_am_pim_Synth	Pyr + SuccCoA + ATP + 2 NADPH + Asp + Glu ==> alKG + Succ + di_am_pim
99	ADPGlc_Synth	G6P + ATP ==> ADPGlc
100	Glucose uptake	==> GLC
101	Glycerol exchange	Glyc ==>
102	<i>glp</i>	Glyc3P ==> ATP + Glyc
103	<i>pfk</i>	F6P + ATP ==> F16P
104	<i>mae</i>	Mal ==> Pyr + NADPH + CO ₂
105	<i>Oxygen uptake</i>	==> O ₂
106	<i>cyc</i>	QuiH ₂ + 0.5 O ₂ ==> 2 H ₂ ex
107	<i>Asnb</i>	2 ATP + Asp + Gln ==> Glu + Asn
108	<i>glbD</i>	alKG + NADPH + Gln ==> 2 Glu
109	<i>Cys</i>	S + ASER ==> Ac + Cys
110	<i>ilvB</i>	2 Pyr ==> CO ₂ + ACLAC
111	<i>ilvC</i>	NADPH + ACLAC ==> DHVAL
112	<i>ilvD</i>	DHVAL ==> OIVAL
113	<i>proB</i>	ATP + Glu ==> GLUP
114	<i>proA</i>	NADPH + GLUP ==> GLUGSAL
115	<i>aroF</i>	PEP + E4P ==> 3DDAH7P
116	<i>aroB</i>	3DDAH7P ==> DQT
117	<i>aroD</i>	DQT <=> DHSK
118	<i>aroE</i>	NADPH + DHSK <=> SME
119	<i>aroL</i>	ATP + SME ==> SME5P
120	<i>aroC</i>	PEP + SME5P ==> 3PSME
121	<i>thrA</i>	ATP + Asp <=> BASP
122	<i>asd</i>	2 NADPH + BASP <=> HSER
123	<i>metL</i>	ATP + HSER ==> PHSER
124	<i>sera</i>	3PG ==> NADH + PHP
125	<i>serC</i>	Glu + PHP ==> alKG + 3PSER

Table S1 (continued)

126	<i>pheA</i>	Chor ==> PHEN
127	<i>pheA2</i>	PHEN ==> CO2 + PHPYR
128	<i>trpDE</i>	Chor + Gln ==> Pyr + Glu + AN
129	<i>trpD</i>	PRPP + AN ==> NPRAN
130	<i>trpC</i>	NPRAN ==> CPAD5P
131	<i>trpC2</i>	CPAD5P ==> CO2 + IGP
132	<i>tyrA</i>	PHEN ==> NADH + CO2 + HPHPYR
133	<i>argA</i>	AcCoA + Glu ==> NAGLU
134	<i>argB</i>	ATP + NAGLU ==> NAGLUYP
135	<i>argC</i>	NADPH + NAGLUYP <=> NAGLUSAL
136	<i>argD</i>	Glu + NAGLUSAL <=> a1KG + NAARON
137	<i>argE</i>	NAARON ==> Ac + ORN
138	<i>carAB</i>	2 ATP + CO2 + Gln ==> Glu + CAP
139	<i>argFI</i>	ORN + CAP <=> CITR
140	<i>argG</i>	2 ATP + Asp + CITR ==> ARGSUCC
141	<i>ilvA</i>	Thr ==> N + OBUT
142	<i>ilvBN</i>	Pyr + OBUT ==> CO2 + ABUT
143	<i>ilvC2</i>	NADPH + ABUT ==> DHMVA
144	<i>ilvD2</i>	DHMVA ==> OMVAL
145	<i>hisG</i>	ATP + PRPP ==> PRBATP
146	<i>hisI</i>	PRBATP ==> PRBAMP
147	<i>hisE</i>	PRBAMP ==> PRFP
148	<i>hisA</i>	PRFP ==> PRLP
149	<i>hisF</i>	Gln + PRLP ==> Glu + DIMGP
150	<i>hisB</i>	DIMGP ==> IMACP
151	<i>hisC</i>	Glu + IMACP ==> a1KG + HISOLP
152	<i>hisB2</i>	HISOLP ==> HISOL
153	<i>metA</i>	SuccCoA + HSER ==> OSLHSER
154	<i>metB</i>	Cys + OSLHSER ==> Succ + LLCT
155	<i>metC</i>	LLCT ==> Pyr + N + HCYS
156	<i>metF</i>	NADH + METTHF ==> MTHF
157	<i>leuA</i>	AcCoA + OIVAL ==> CBHCAP
158	<i>leuCD</i>	CBHCAP <=> IPPMAL
159	<i>leuB</i>	IPPMAL ==> NADH + CO2 + OICAP

Metabolites list:

2PG	2-Phosphoglycerate
3DDAH7P	3-Deoxy-d-arabino heptulosonate-7-phosphate
3PG	3-Phosphoglycerate
3PSER	3-Phosphoserine
3PSME	3-Phosphate-shikimate
ABUT	2-Aceto-2-hydroxy butyrate
Ac	Acetate
AcCoA	Acetyl-CoA
ACLAC	Acetolactate
AcP	Acetyl phosphate
Adh	Acetaldehyde
ADPGlc	ADPglucose

Ala	Alanine
alKG	alpha-Ketoglutarate
AN	Antranilate
Arg	Arginine
ARGSUCC	L-Arginio succinate
ASER	O-Acetylserine
Asn	Asparagine
Asp	Aspartate
ATP	Adenosintriphosphate
avg_FS	average fatty acid
BASP	b-Aspartyl phosphate
Biomass	Biomass
C14_0_FS	C_14:0_Fatty_acid
CAP	Carbamoyl phosphate
CBHCAP	3-Carboxy-3-hydroxy-isocaproate
CDPEth	CDP ethanolamine
Chor	Chorismate
Cit	Citrate
CITR	L-Citrulline
CMP_KDO	CMP-3-deoxy-D-manno-octulosonate
CO2	Carbon dioxide
CPAD5P	1-O-Carboxyphenylamino 1-deoxyribulose-5-phosphate
Cys	Cysteine
dATP	ATP for DNA synthesis
dCTP	CTP for DNA synthesis
dGTP	GTP for DNA synthesis
DHAP	Dihydroxyacetone phosphate
DHMVA	2,3-Dihydroxy-3-methyl-valerate
DHSK	Dehydroshikimate
DHVAL	Dihydroxy-isovalerate
di_am_pim	Diaminopimelate
DIMGP	D-Erythro imidazoleglycerol-phosphate
DPG	Diphosphoglycerate
DQT	3-Dehydroquinate
dTTP	TTP for DNA synthesis
E4P	D-Erythrose 4-phosphate
Eth	Ethanol
F16P	Fructose 1,6-bisphosphate
F6P	Fructose 6-phosphate
Form	Formate
Fum	Fumarate
G3P	Glyceraldehyde 3-phosphate
G6P	Glucose 6-phosphate
GLC	Glucose
Gln	Glutamine
Glu	Glutamate
GLUGSAL	L-Glutamate g-semialdehyde
GLUP	Glutamyl phosphate
Gly	Glycine
Glyc	Glycerol
Glyc3P	Glycerol 3-phosphate
Glyox	Glyoxylate
H_ex	External Hydrogen
HCYS	Homocysteine
His	Histidine
HISOL	Histidinol

HISOLP	L-Histidinol-phosphate
HPPYR	para-Hydroxy phenyl pyruvate
HSER	Homoserine
ICit	Isocitrate
IGP	Indole glycerol phosphate
Ile	Isoleucine
IMACP	Imidazole acetyl-phosphate
IPPMAL	3-Isopropylmalate
KetoPGLuc	2-keto-3-deoxy-D-gluconate 6-phosphate
Lac	Lactate
Leu	Leucine
LLCT	L-Cystathionine
Lys	Lysine
Mal	Malate
Met	Methionine
METTHF	5,10-Methylene tetrahydrofolate
MTHF	Methylen-Tetrahydrofolate
N	Nitrogen(NH ₄)
NAARON	N-a-Acetyl ornithine
NADH	Nicotinamide adenine dinucleotide - reduced
NADPH	Nicotinamide adenine dinucleotide phosphate - reduced
NAGLU	N-Acetyl glutamate
NAGLUSAL	N-Acetyl glutamate semialdehyde
NAGLUYP	N-Acetyl glutamyl -phosphate
NDPHep	NDP Heptose
NPRAN	N-5-phosphoribosyl-antranilate
O ₂	Oxygen
OBUT	Oxobutyrate or 2-ketobutyrate
OH_myristic_ac	OH myristic Acid
OICAP	2-Oxoisocaproate
OIVAL	Oxoisovalerate
OMVAL	Oxomethylvalerate
ORN	Ornithine
OSLHSER	O-Succinyl-l-homoserine
OxA	Oxaloacetate
PEP	Phosphoenolpyruvate
PGLac	6-Phospho-Gluconolactone
PGLuc	6-Phospho-Gluconate
Phe	Phenylalanine
PHEN	Prephenate
PHP	3-Phosphohydroxypyruvate
PHPYR	Phenyl pyruvate
PHSER	O-Phospho-l-homoserine
PRBAMP	Phosphoribosyl -AMP
PRBATP	Phosphoribosyl-ATP
PRFP	Phosphoribosyl-formimino-AICAR-phosphate
PRLP	Phosphoribulosyl- formimino-AICAR-phosphate
Pro	Proline
PRPP	5-Phospho-alpha-D-ribose 1-diphosphate
Pyr	Pyruvate
QuiH ₂	Ubichinon_red
R5P	Ribose 5-phosphate
rATP	ATP for RNA synthesis
rCTP	CTP for RNA synthesis
rGTP	GTP for RNA synthesis
R15P	Ribulose 5-phosphate

rUTP	UTP for RNA synthesis
S	Sulfur(SO4)
S7P	Sedoheptulose 7-phosphate
Ser	Serine
SME	Shikimate
SME5P	Shikimate-5-phosphate
Succ	Succinate
SuccCoA	Succinyl-CoA
TDPGlc	TDP-glucosamine
Thr	Threonine
Trp	Tryptophan
Tyr	Tyrosine
UDP_NAG	UDP acetylglucosamine
UDP_NAM	UDP N-acetylmuramic acid
UDPGlc	UDP glucose
Val	Valine
X5P	Xylolose-5-Phosphate

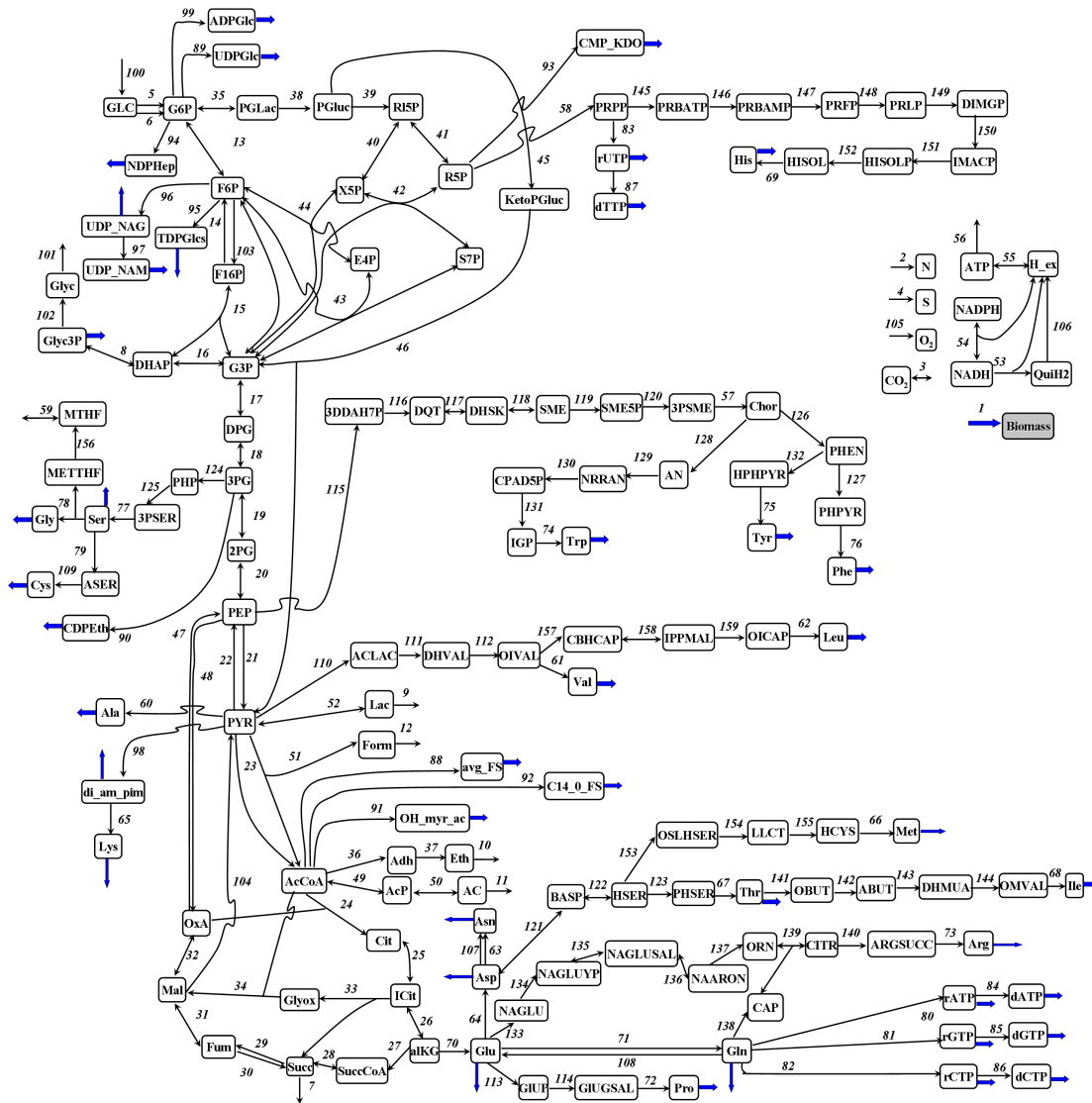


Figure S1 – Metabolic network map for *E. coli*

Details of the metabolic reactions and metabolites of *E. coli* are shown in **Table S1**.

Table S2 Metabolic reaction models for *S. Cerevisiae*

Gene	Enzyme	Reaction
1Put1m	Proline oxidase (NAD)	NADMIT + PRO ==> NADHMIT + 1PYR5C
AAT2	Tyrosine transaminase	AKG + TYR <=> GLU + 34HPP
AATA	2-aminoadipate transaminase	GLU + 2OXOADP <=> AKG + L2AADP
ACO	Aconitate synthetase	CIT <=> ISOCIT
ACS	Acetyl-coenzyme A synthetase	2 ATP + AC ==> ACCOACYT + 2 ADP
ACXT	Acetate secretion	AC ==> ACXT
ADH14G	Acetaldehyde dehydrogenase	ACAL + NADHCYT ==> NADCYT + ETOH
ALD4	Cytosolic aldehyde dehydrogenase (mitochondrial)	ACAL + NADMIT ==> NADHMIT + AC
ALD6	Cytosolic aldehyde dehydrogenase	ACAL + NADPCYT ==> NADPHCYT + AC
Alt2m	L-alanine transaminase, mitochondrial	AKG + ALA <=> PYR + GLU
Aro1	Shikimate kinase	ATP + SKM ==> ADP + SKM5P
Aro1b	shikimate dehydrogenase	NADPHCYT + 3DHSK ==> NADPCYT + SKM
Aro1c	3-phosphoshikimate 1-carboxyvinyltransferase	PEP + SKM5P ==> 3PSME
Aro1d	3-dehydroquinate dehydratase	3DHQ ==> 3DHSK
Aro1e	3-dehydroquinate synthase	2DDA7P ==> 3DHQ
Aro2	Chorismate synthase	3PSME ==> CHOR
Aro3	3-deoxy-D-arabino-heptulosonate 7-phosphate synthetase	PEP + E4P ==> 2DDA7P
Aro7	Chorismate mutase	CHOR ==> PPHN
Aro9	Phenylalanine transaminase	AKG + PHE <=> GLU + PHPYR
ART2	Aspartate transaminase	AKG + ASP <=> OAC + GLU
ASN123	Asparagine synthase (glutamine-hydrolysing)	2 ATP + ASP + GLN ==> 2 ADP + GLU + ASN
ASP1	L-asparaginase	ASN ==> ASP
Bat	Valine transaminase	AKG + VAL <=> GLU + 3MOB
Bat2b	2-Oxo-4-methyl-3-carboxypentanoate decarboxylation	3C4MOP ==> CO2 + 4MOP
Bat2c	Leucine transaminase	AKG + LEU <=> GLU + 4MOP
BIOMX	Biomass formation	435 P3G + 1639 NADPHMIT + 24118 ATP + 300 RIB5P + 128 PEP + 2104 ACCOACYT + 5552 NADPHCYT + 528 AKG + 601 OAC + 64 E4P + 14 ACCOAMIT + 314 NADMIT + 100 GOH3P + 833 NADCYT + 2500 GLUC6P + 219 PYR + 297 ASP + 302 GLU + 105 GLN + 102 ASN + 459 ALA + 165 PRO + 185 SER + 286 LYS + 102 TYR + 134 PHE + 265 VAL + 296 LEU ==> 314 NADHMIT + 1639 NADPMIT + 833 NADHCYT + 24118 ADP + 5552 NADPCYT + 1000 BIOM
CAT	Carnitine O-acetyltransferase, Carnitine shuttle	ACCOACYT ==> ACCOAMIT
CIT	Citrite synthase (mitochondrial)	OAC + ACCOAMIT ==> CIT
CIT2	Cytosolic Citrite synthase	ACCOACYT + OAC ==> CIT
CO2XT	CO2 secretion	CO2 ==> CO2XT

Table S2 (continued)

ENO	Enolase	$P2G \rightleftharpoons PEP$
FADHX	Electronic chain: Reoxidation of FADH2 P/O ratio 1.2	$20 FADH2 + 24 ADP + 10 O2 \implies 24 ATP + 20 FAD$
FBA	Fructose-1,6 phosphate aldorase	$FRUCDP \rightleftharpoons DHAP + GA3P$
FBP	Fructose-1,6-bisphosphatase	$FRUCDP \implies FRUC6P$
FUM1	Fumarate hydratase	$FUM \rightleftharpoons MAL$
Gad1	Glutamate Decarboxylase	$GLU \implies CO2 + 4ABUT$
Gdh1	Glutamate dehydrogenase (NADP)	$NADPCYT + GLU \rightleftharpoons NADPHCYT + AKG$
Gdh2	Glutamate dehydrogenase (NAD)	$NADCYT + GLU \implies AKG + NADHCYT$
GLK	Glucokinase	$ATP + GLUC \implies GLUC6P + ADP$
Gln1	Glutamine synthetase	$ATP + GLU \implies ADP + GLN$
Glt1	Glutamate synthase (NADH2)	$AKG + NADHCYT + GLN \implies NADCYT + 2 GLU$
GLUN	Glutaminase	$GLN \implies GLU$
GND	6-Phosphogluconate dehydrogenase	$P6G + NADPCYT \implies RIBL5P + NADPHCYT + CO2$
GPD	Glycerol -3-phosphate	$DHAP + NADHCYT \implies GOH3P + NADCYT$
GPM	Phosphoglycerate mutase	$P3G \rightleftharpoons P2G$
GPP	Glycerol phosphatase	$GOH3P \implies GOH$
HCITSm	Homocitrate synthase, mitochondrial	$AKG + ACCOAMIT \implies HCIT$
ICL	Isocitrate lyase	$ISOCIT \implies GLYO + SUC$
IDH	Isocitrate dehydrogenase NAD	$ISOCIT + NADMIT \implies NADHMIT + AKG + CO2$
IDP1	Isocitrate dehydrogenase (NADPmit-dependent)	$NADPMIT + ISOCIT \implies NADPHMIT + AKG + CO2$
IDP2	Isocitrate dehydrogenase (NADcyt-dependent)	$ISOCIT + NADPCYT \implies NADPHCYT + AKG + CO2$
Ilv26m	Acetolactate synthase, mitochondrial	$2 PYR \implies CO2 + ALACS$
Ilv3m	Dihydroxy-acid dehydratase (2,3-dihydroxy-3-methylbutanoate), mitochondrial	$23DHMB \implies 3MOB$
Ilv5m	Acetohydroxy acid isomeroreductase, mitochondrial	$NADPHMIT + ALACS \implies NADPMIT + 23DHMB$
KGD	Alpha-ketoglutarate dehydrogenase	$AKG + NADMIT \implies NADHMIT + CO2 + SUCCOA$
Leu1	2-isopropylmalate hydratase	$2IPPM \rightleftharpoons 3C3HMP$
Leu1b	3-isopropylmalate dehydratase	$3C2HMP \rightleftharpoons 2IPPM$
Leu2	3-isopropylmalate dehydrogenase	$NADCYT + 3C2HMP \implies NADHCYT + 3C4MOP$
Leu4	2-isopropylmalate synthase, mitochondrial	$ACCOACYT + 3MOB \implies 3C3HMP$
LSC	Succinate-CoA ligase	$ADP + SUCCOA \rightleftharpoons ATP + SUC$
Lys1	Saccharopine dehydrogenase (NAD, L-lysine forming)	$NADCYT + SACCRP \rightleftharpoons AKG + NADHCYT + LYS$
Lys12m	Homoisocitrate dehydrogenase, mitochondrial	$NADMIT + HICIT \rightleftharpoons NADHMIT + OXAG$
Lys25	L-aminoadipate-semialdehyde dehydrogenase (NADPH)	$2 ATP + NADPHCYT + L2AADP \implies 2 ADP + NADPCYT + L2AADP6SA$
Lys4m	Homoacontinate hydratase, mitochondrial	$B124TC \rightleftharpoons HICIT$

Table S2 (continued)

Lys9	Saccharopine dehydrogenase (NADP, L-glutamate forming)	NADPHCYT + GLU + L2AADP6SA \rightleftharpoons NADPCYT + SACCRP
MAE	Malic enzyme	MAL + NADPMIT \implies NADPHMIT + CO ₂ + PYR
MAINT	maintenance	ATP \implies ADP
MCITDm	2-methylcitrate dehydratase, mitochondrial	HCIT \rightleftharpoons B124TC
MDH	Malase dehydrrogenase	MAL + NADMIT \rightleftharpoons NADHMIT + OAC
MDH2	Malase dehydrrogenase (NADcyt dependent)	MAL + NADCYT \rightleftharpoons OAC + NADHCYT
MLS	Malase synthase	GLYO + ACCOACYT \implies MAL
NADHX	Electronic chain: Reoxidation of NADH P/O ratio 1.2	20 NADHMIT + 24 ADP + 10 O ₂ \implies 24 ATP + 20 NADMIT
O2XT	Oxygen input	O2XT \implies O ₂
OSM	Fumarate reductase	FADH ₂ + FUM \implies FAD + SUC
OXAGm	non-enzymatic reaction, mitochondrial	OXAG \rightleftharpoons CO ₂ + 2OXOADP
P5CDm	1-pyrroline-5-carboxylate dehydrogenase	NADMIT + 1PYR5C \implies NADHMIT + GLU
PCK	Phosphoenolpyruvate carboxykinase	ATP + OAC \implies PEP + CO ₂ + ADP
PDA	Pyruvate dehydrogenase	NADMIT + PYR \implies NADHMIT + ACCOAMIT + CO ₂
PDC	Pyruvate decarboxylase	PYR \implies ACAL + CO ₂
PFK	Phosphofructokinase	ATP + FRUC6P \implies ADP + FRUCDP
PGI	Glucose-6-phosphate isomerase	GLUC6P \rightleftharpoons FRUC6P
PGK	3-Phosphoglycerate kinase	P13G + ADP \rightleftharpoons P3G + ATP
Pha2	Prephenate dehydratase	PPHN \implies CO ₂ + PHPYR
Pro1	Glutamate 5-kinase	ATP + GLU \implies ADP + GLU5P
Pro2	Glutamate-5-semialdehyde dehydrogenase	NADPHCYT + GLU5P \implies NADPCYT + GLU5SA
Pro2b	L-glutamate 5-semialdehyde dehydratase	GLU5SA \rightleftharpoons 1PYR5C
Pro3	Pyrroline-5-carboxylate reductase	NADPHCYT + 1PYR5C \implies NADPCYT + PRO
PYC	Pyruvate carboxylase	ATP + CO ₂ + PYR \implies OAC + ADP
PYK	Pyruvate kinase	PEP + ADP \implies ATP + PYR
RKI	Ribose 5-phosphate isomerase	RIBL5P \rightleftharpoons RIB5P
RPE	Ribulose-phosphate 3-epimerase	RIBL5P \rightleftharpoons XYL5P
SDH	Succinate dehydrogenase	FAD + SUC \implies FADH ₂ + FUM
Ser1	Phosphoserine transaminase	GLU + 3PHP \implies AKG + PSER
Ser2	Phosphoserine phosphatase (L-serine)	PSER \implies SER
Ser3	Phosphoglycerate dehydrogenase	P3G + NADCYT \implies NADHCYT + 3PHP
SHUTTLEX	Reoxidation of cytosolic NADH to mitochondrial NADH	NADHCYT + NADMIT \implies NADHMIT + NADCYT
SOL	6-phosphoglucono-Lactonase	G15L \implies P6G
TAL	Transaldolase	SED7P + GA3P \rightleftharpoons E4P + FRUC6P
TDH	Glyceraldehyde-3-phosphate dehydrogenase	GA3P + NADCYT \rightleftharpoons NADHCYT + P13G
TKI	Transaldolase II	E4P + XYL5P \rightleftharpoons GA3P + FRUC6P
TKL	Transketolase	RIB5P + XYL5P \rightleftharpoons SED7P + GA3P

Table S2 (continued)

TPI	Triosephosphate isomerase	DHAP \rightleftharpoons GA3P
TYR1	Prephenate dehydrogenase (NADP)	NADPCYT + PPHN \implies NADPHCYT + CO ₂ + 34HPP
Uga1	4-aminobutyrate transaminase	AKG + 4ABUT \implies GLU + SUCSAL
Uga2	Succinate-semialdehyde dehydrogenase (NADP)	NADPCYT + SUCSAL \implies NADPHCYT + SUC
ZWF	Glucose-6-phosphate dehydrogenase	GLUC6P + NADPCYT \implies G15L + NADPHCYT

\rightleftharpoons reversible reaction;

\implies irreversible reaction;

Metabolites:

1PYR5C	(S)-1-Pyrroline-5-carboxylate
23DHMB	(R)-2,3-Dihydroxy-3-methylbutanoate
2DDA7P	2-Dehydro-3-deoxy-D-arabino-heptonate 7-phosphate
2IPPM	2-Isopropylmaleate
2OXOADP	2-Oxoadipate
34HPP	3-(4-Hydroxyphenyl)pyruvate
3C2HMP	3-Carboxy-2-hydroxy-4-methylpentanoate
3C3HMP	3-Carboxy-3-hydroxy-4-methylpentanoate
3C4MOP	3-Carboxy-4-methyl-2-oxopentanoate
3DHQ	3-Dehydroquininate
3DHSK	3-Dehydroshikimate
3MOB	3-Methyl-2-oxobutanoate
3PHP	3-Phosphonooxypyruvate
3PSER	3-Phosphoserine
3PSME	5-O-(1-Carboxyvinyl)-3-phosphoshikimate
4ABUT	4-Aminobutanoate
4MOP	4-Methyl-2-oxopentanoate
AC	Acetate
ACAL	Acetaldehyde
ACCOACYT	Acetyl-coenzyme A (cytosolic)
ACCOAMIT	Acetyl-coenzyme A (mitochondria)
ACXT	External acetate
ADP	Adenosine-5'-diphosphate
AKG	Alpha-ketoglutarate
ALA	Alanine
ALACS	(S)-2-Acetolactate
ASN	Asparagine
ASP	Aspartate
ATP	adenosine triphosphate
B124TC	But-1-ene-1,2,4-tricarboxylate
BIOM	Biomass
CHOR	Chorismate
CIT	Citrate
CO2	Carbon dioxide (External)
CO2XT	External Carbon dioxide
DHAP	Dihydroxyacetone phosphate
E4P	Erthrose-4-phosphate
ETOH	External ethanol
FAD	Flavin adenine dinucleotide
FADH2	Flavin adenine dinucleotide, reduced
FRUC6P	Fructose-6-phosphate
FRUCDP	D-Fructose 1,6-bisphosphate

FUM	Fumarate
G15L	6-phospho-D-glucono-1,5-lactone
GA3P	Glyceraldehyde-3-phosphate
GLU	Glutamate
GLU5P	alpha-D-Glutamyl phosphate
GLU5SA	L-Glutamate 5-semialdehyde
GLUC	Glucose
GLUC6P	Glucose-6-phosphate
GLN	Glutamine
GLYO	Glyoxylate
GOH	Glycerol
GOH3P	Glycerol-3-phosphate
HCIT	2-Hydroxybutane-1,2,4-tricarboxylate
ISOCIT	Isocitrate
L2AADP	L-2-Amino adipate
L2AADP6SA	L-2-Amino adipate 6-semialdehyde
LEU	Leucine
MAL	Malate
NADCYT	Nicotinamide adenine dinucleotide (cytosolic)
NADHCYT	Nicotinamide adenine dinucleotide - reduced (cytosolic)
NADHMIT	Nicotinamide adenine dinucleotide - reduced (mitochondrial)
NADMIT	Nicotinamide adenine dinucleotide (mitochondrial)
NADPCYT	Nicotinamide adenine dinucleotide phosphate (cytosolic)
NADPHCYT	Nicotinamide adenine dinucleotide phosphate - reduced (cytosolic)
NADPHMIT	Nicotinamide adenine dinucleotide phosphate - reduced
NADPMIT	Nicotinamide adenine dinucleotide phosphate (mitochondrial)
O2	Oxygen
O2XT	External Oxygen
OAC	Oxaloacetate
OXAG	Oxaloglutarate
P13G	3-Phospho-D-glyceroyl phosphate
P2G	D-Glycerate 2-phosphate
P3G	D-Glycerate 3-phosphate
P6G	6-Phospho-D-gluconate
PEP	Phosphoenolpyruvate
PHE	Phenylalanine
PHPYR	Phenylpyruvate
PPHN	Prephenate
PRO	Proline
PYR	Pyruvate
RIB5P	Ribose-5-phosphate
RIBL5P	Ribulose-5-phosphate
SACCRP	N6-(L-1,3-Dicarboxypropyl)-L-lysine
SED7P	Sedoheptulose-7-phosphate
SER	Serine
SKM	Shikimate
SKM5P	Shikimate 3-phosphate
SUC	Succinate
SUCSAL	Succinate semialdehyde
SUCCOA	Succinyl-coenzyme A
TYR	Tyrosine
VAL	valine
XYL5P	Xylulose-5-phosphate

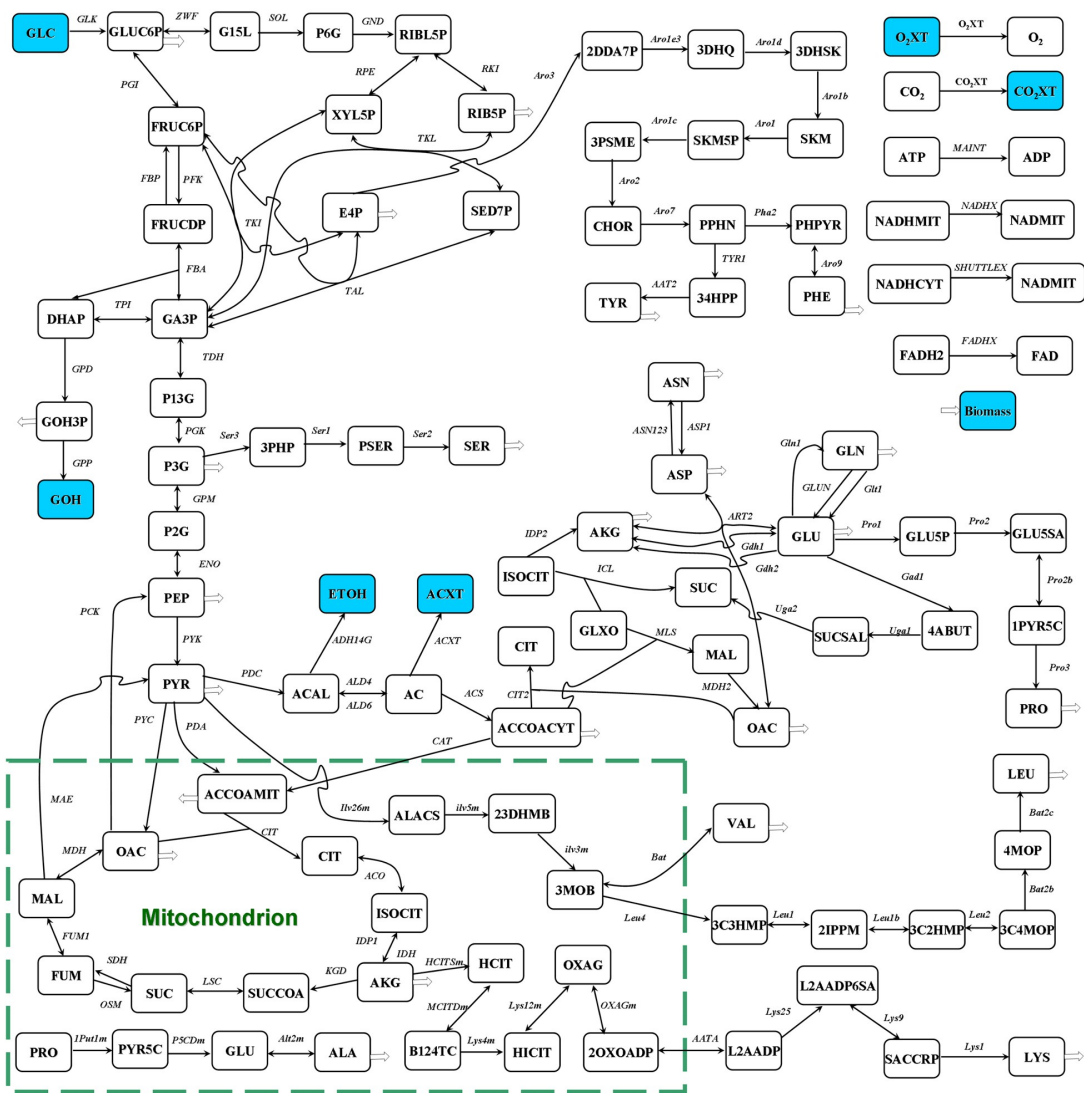


Figure S2 Metabolic network model for *S. cerevisiae* (Blue metabolites are external ones)