

Table S1. Reactions in the yeast meiosis-specific metabolic network.

Reaction	Metabolite ^{&}	Enzyme*	Objective function [^]	Pathway	Reference
<i>Reactions occurring in the cytoplasm</i>					
V1: Coenzyme A + Acetate + ATP => Acetyl-CoA + Pyrophosphate + AMP	Acetate, Acetyl-CoA	<i>ACS2</i>	Net ATP production (-1), ATP consumption (1)		[1,2]
V2: Oxaloacetate + Acetyl-CoA + H ₂ O => Citrate + Coenzyme A	Oxaloacetate, Acetyl-CoA, Citrate	<i>CIT2</i>			[1,2,3,4]
V3: Citrate + H ₂ O => Isocitrate	Citrate, Isocitrate	<i>ACO1</i>			[1,2,3]
V4: Isocitrate => Glyoxylate + Succinate	Isocitrate, Glyoxylate, Succinate	<i>ICL1</i>		Glyoxylate	[1,2,3]
V5: Glyoxylate + Acetyl-CoA => Malate	Glyoxylate, Acetyl-CoA, Malate	<i>MLS1</i>		Glyoxylate	[1,2,3]
V6: Malate + NAD => Oxaloacetate + NADH + H ⁺	Malate, Oxaloacetate	<i>MDH2</i>	Net ATP production (3), ATP production (3)		[1,2,3]
V7: Oxaloacetate + GTP => CO ₂ + Phosphoenolpyruvate + GDP	Oxaloacetate, Phosphoenolpyruvate	<i>PCK1</i>	Net ATP production (-1), ATP consumption (1)	Gluconeogenesis	[2,5]
V8: Phosphoenolpyruvate + H ₂ O => 2-Phosphoglycerate	Phosphoenolpyruvate, 2-Phosphoglycerate	<i>ENO1</i>		Gluconeogenesis	[2,3]
V9: 2-Phosphoglycerate => 3-Phosphoglycerate	2-Phosphoglycerate, 3-Phosphoglycerate	<i>GPM1</i>		Gluconeogenesis	[2,3]
V10: 3-Phosphoglycerate + ATP => 1,3-BIP Glycerate + ADP	3-Phosphoglycerate, 1,3-BIP Glycerate	<i>PGK1</i>	Net ATP production (-1), ATP consumption	Gluconeogenesis	[2,3,5]

		(1)		
V11: 1,3-BIP Glycerate + NADH + H+ => Glyceraldehyde-3-Phosphate + phosphate + NAD	1,3-BIP Glycerate, Glyceraldehyde-3-Phosphate	<i>TDH1</i> <i>TDH2</i> <i>TDH3</i> (or)	Net ATP production (-3), ATP consumption (3)	Gluconeogenesis [2,3,6]
V12: Glyceraldehyde-3-Phosphate => Dihydroxy-acetone-phosphate	Glyceraldehyde-3-Phosphate, Dihydroxy-acetone-phosphate	<i>TPI1</i>		Gluconeogenesis [2,3,6]
V13: Dihydroxy-Acetone-phosphate + Glyceraldehyde-3-phosphate => Fructose-1,6-bisphosphate	Dihydroxy-Acetone-phosphate, Glyceraldehyde-3-phosphate, Fructose-1,6-bisphosphate	<i>FBA1</i>		Gluconeogenesis [2,3,6]
V14: Fructose-1,6-bisphosphate + H2O => Fructose-6-phosphate + phosphate	Fructose-1,6-bisphosphate, Fructose-6-phosphate	<i>FBP1</i>		Gluconeogenesis [2,3,6]
V15: Fructose-6-phosphate => Glucose-6-phosphate	Fructose-6-phosphate, Glucose-6-phosphate	<i>PGI1</i>		Gluconeogenesis [2,3,6]
V16: Glucose-6-phosphate => Glucose-1-phosphate	Glucose-6-phosphate, Glucose-1-phosphate	<i>PGM2</i>		Gluconeogenesis [2,3]
V17: UDP-D-glucose + Glucose-6-phosphate => UDP + α,α -trehalose 6-phosphate	Glucose-6-phosphate	<i>TPS1</i> <i>TPS2</i> <i>TPS3</i> <i>TLS1</i> (and)	Carbohydrate synthesis	Gluconeogenesis [2,7,8]
α,α -trehalose 6-phosphate + H2O => Trehalose + phosphate				
V18: Glucose-1-phosphate + UTP => UDP-Glucose + 2 Pi	Glucose-1-phosphate	<i>UGPI</i>	Carbohydrate synthesis	Gluconeogenesis [2,8,9,10]
Glycogen _(n residues) + UDP-Glucose => Glycogen _(n+1 residues) + UDP				
V19: a Glycogen + Phosphate => a Glycogen + Glucose-1-phosphate	Glucose-1-phosphate	<i>GPH1</i>	Carbohydrate breakdown	Glycogenolysis [2,11,12]

V20: Glucose-1-phosphate => Glucose-6-phosphate	Glucose-1-phosphate, Glucose-6-phosphate	<i>PGM2</i>		Glycogenolysis	[1,2,3]
V21: Glucose-6-phosphate => Fructose-6-phosphate	Glucose-6-phosphate, Fructose-6-phosphate	<i>PGII</i>		Glycogenolysis	[2,3,6]
V22: Fructose-6-phosphate +ATP => Fructose-1,6- biphosphate + ADP + H ⁺	Fructose-6-phosphate , Fructose-1,6- biphosphate	<i>PFK1</i> <i>PFK2</i> (and)	Net ATP production (- 1), ATP consumption (1)	Glycogenolysis	[2,3,6]
V23: Fructose-1,6- biphosphate => Dihydroxy-Acetone- phosphate + Glyceraldehyde-3- phosphate	Fructose-1,6- biphosphate, Dihydroxy-Acetone- phosphate, Glyceraldehyde-3- phosphate	<i>FBA1</i>		Glycogenolysis	[2,3,6]
V24: Dihydroxy-Acetone- phosphate => Glyceraldehyde-3- phosphate	Dihydroxy-Acetone- phosphate, Glyceraldehyde-3- phosphate	<i>TPI1</i>		Glycogenolysis	[2,3,6]
V25: Glyceraldehyde-3- phosphate + Phosphate + NAD => 1,3-BIP Glycerate + NADH + H ⁺	Glyceraldehyde-3- phosphate, 1,3-BIP Glycerate	<i>TDH1</i> <i>TDH2</i> <i>TDH3</i> (or)	Net ATP production (3), ATP production (3)	Glycogenolysis	[2,3,6]
V26: 1,3-BIP Glycerate + ADP => 3- Phosphoglycerate + ATP	1,3-BIP Glycerate, 3-Phosphoglycerate	<i>PGK1</i>	Net ATP production (1), ATP production (1)	Glycogenolysis	[2,3,6]
V27: 3-Phosphoglycerate => 2-Phosphoglycerate	3-Phosphoglycerate, 2- Phosphoglycerate	<i>GPM1</i>		Glycogenolysis	[2,3,6]
V28: 2-Phosphoglycerate => Phosphoenolpyruvate + H2O	2-Phosphoglycerate, Phosphoenolpyruvate	<i>ENO1</i>		Glycogenolysis	[2,3,6]
V29: Phosphoenolpyruvate + ADP + H ⁺ => Pyruvate + ATP	Phosphoenolpyruvate, Pyruvate	<i>PYK2</i>	Net ATP production (1), ATP production (1)	Glycogenolysis	[2,3,6]

V30: HCO ₃ ⁻ + Pyruvate + ATP => Phosphate + Oxaloacetate + ADP	Pyruvate, Oxaloacetate	<i>PYC1</i> <i>PYC2</i> (or)	Net ATP production (-1), ATP consumption (1)	Glycogenolysis	[1,2]
V31: Fumarate + H ₂ O => Malate	Fumarate, Malate	<i>FUM1</i>			[2,3,5]
V32: Isocitrate + NADP => Oxoglutarate + CO ₂ + NADPH	Isocitrate, Oxoglutarate	<i>IDP2</i>	Net ATP production (3), ATP production (3)	Glutamate	[2,3]
V33: Oxoglutarate + NH ₃ + NADPH + H ⁺ => L-Glutamate + H ₂ O + NADP	Oxoglutarate	<i>GDH3</i>	Net ATP production (-3), ATP consumption (3), Glutamate synthesis	Glutamate	[2,3,8]
Reactions occurring in the mitochondria					
V34: Coenzyme A [m] + NAD + Pyruvate [m] => Acetyl-CoA [m] + CO ₂ + NADH	Pyruvate [m], Acetyl-CoA [m]	<i>PDB1</i> <i>PDA1</i> (and)		Glycogenolysis	[2,3,6]
V35: Oxaloacetate [m] + Acetyl-CoA [m] + H ₂ O => Citrate [m] + Coenzyme A [m]	Oxaloacetate [m], Acetyl-CoA [m], Citrate [m]	<i>CIT1</i>		TCA	[1,2,3,4]
V36: Citrate [m] + H ₂ O => Isocitrate [m]	Citrate [m], Isocitrate [m]	<i>ACO1</i>		TCA	[1,2,3]
V37: NAD + Isocitrate [m] => Oxoglutarate [m] + NADH + CO ₂	Isocitrate [m], Oxoglutarate [m]	<i>IDH1</i> <i>IDH2</i> (and)	Net ATP production (3), ATP production (3)	TCA	[1,2,3]
V38: Oxoglutarate [m] + Coenzyme A [m] + NAD => Succinyl-CoA [m] + CO ₂ + NADH	Oxoglutarate [m], Succinyl-CoA [m]	<i>LPD1</i> <i>KGD1</i> <i>KGD2</i> (and)	Net ATP production (3), ATP	TCA	[2,3,5]

				production (3)	
V39: Succinyl-CoA [m] + GDP + Phosphate <=> Succinate [m] + Coenzyme A [m] + GTP	Succinyl-CoA [m], Succinate [m]	<i>LSC1</i> <i>LSC2</i> (and)	Net ATP production (1), ATP production (1)	TCA	[2,3,5]
V40: Succinate [m] + FAD => Fumarate [m] + FADH2	Succinate [m], Fumarate [m]	<i>SDH1</i> <i>SDH2</i> <i>SDH3</i> <i>SDH4</i> (and)	Net ATP production (2), ATP production (2)	TCA	[1,2,3]
V41: Fumarate [m] + H2O => Malate [m]	Fumarate [m], Malate [m]	<i>FUM1</i>		TCA	[1,2,3]
V42: Malate [m] + NAD => Oxaloacetate [m] + NADH + H ⁺	Malate [m], Oxaloacetate [m]	<i>MDH1</i>	Net ATP production (3), ATP production (3)	TCA	[1,2,3]
<i>Transport reactions</i>					
V43: Acetate [e] => Acetate	Acetate	<i>ADY2</i>	Acetate uptake		[2,13,14]
V44: Acetyl-CoA + Carnitine => CoA + O-Acetylcarnitine	Acetyl-CoA, Acetyl-CoA [m]	<i>CAT2</i> <i>YAT1</i> <i>YAT2</i> (avg)			[2,15]
O-Acetylcarnitine + CoA => Acetyl-CoA [m] + Carnitine ^s					
V45: Oxaloacetate [m] => Oxaloacetate	Oxaloacetate [m], Oxaloacetate	<i>OAC1</i>			[2,16,17]
V46: Oxaloacetate => Oxaloacetate [m]	Oxaloacetate, Oxaloacetate [m]	<i>OAC1</i>			[2,16,17]
V47: Citrate => Citrate [m]	Citrate, Citrate [m]	<i>CTP1</i>			[2,18]
V48: Isocitrate [m] => Isocitrate	Isocitrate, Isocitrate [m]	<i>CTP1</i>			[2,18]
V49: Fumarate [m] + Succinate => Fumarate + Succinate [m] [#]	Fumarate [m], Succinate, Fumarate, Succinate [m]	<i>SFC1</i>			[2,17,19]

V50: Succinate + Phosphate [m] => Succinate [m] + Phosphate	Succinate [m], Succinate	<i>DIC1</i>			[2,17,19]
V51: Oxoglutarate [m] => Oxoglutarate	Oxoglutarate [m], Oxoglutarate	<i>ODC1</i>		Glutamate	[2,20]
V52: Pyruvate => Pyruvate [m]	Pyruvate, Pyruvate [m]	<i>YIA6</i>		Glycogenolysis	[2,21,22]
Reactions that produce nucleotides, amino acids, and lipids					
V53: Glucose-6-phosphate + NADP => D-6-phosphoglucono-δ-lactone + NADPH + H ⁺	Glucose-6-phosphate	<i>ZWF1</i>	Net ATP production (3), ATP production (3), Nucleotide synthesis	Nucleotide	[2,23,24]
V54: 3-Phosphoglycerate + NAD ⁺ => 3-Phosphohydroxypyruvate + H + NADH	3-Phosphoglycerate	<i>SER3</i> <i>SER33</i> (or)	Net ATP production (3), ATP production (3), Amino acid synthesis	Amino acid	[2,23,24]
V55: Oxaloacetate + L-glutamate => 2-oxoglutarate + L-aspartate	Oxaloacetate	<i>AAT2</i>	Amino acid synthesis	Amino acid	[2,23,24]
V56: Pyruvate [m] + NADPH + H ⁺ + glutamate => CO ₂ + NADP ⁺ + H ₂ O + Alpha-ketoglutarate + valine	Pyruvate [m]	<i>ILV2</i> <i>ILV6</i> (and)	Amino acid synthesis	Amino acid	[2,23,24]
V57: Oxaloacetate [m] + L-Glutamate => 2-Oxoglutarate + L-Aspartate	Oxaloacetate [m]	<i>AAT1</i>	Amino acid synthesis	Amino acid	[2,23,24]
V58: Phosphoenolpyruvate + Skm5p => 3Psm + phosphate	Phosphoenolpyruvate	<i>ARO1</i>	Amino acid synthesis	Amino acid	[2,23,24]
V59: BCCP-biotin-CO ₂ + Acetyl-CoA + ATP => Malonyl-CoA + BCCP-	Acetyl-CoA	<i>ACC1</i>	Net ATP production (-1),	Lipid	[2,23,24]

biotin + phosphate + ADP			ATP consumption (1), Lipid synthesis		
V60: BCCP-biotin-CO ₂ + Acetyl-CoA [m] + ATP => Malonyl-CoA + BCCP-biotin + phosphate + ADP	Acetyl-CoA [m]	<i>HFAI</i>	Net ATP production (-1), ATP consumption (1), Lipid synthesis	Lipid	[2,23,24]
V61: Dihydroxy-Acetone-phosphate + NADH => Glycerol 3 P + NAD	Dihydroxy-Acetone-phosphate	<i>GPD1</i>	Net ATP production (-3), ATP consumption (3), Lipid synthesis	Lipid	[2,23,24]
V62: Glucose-6-phosphate => Mi1p-D	Glucose-6-phosphate	<i>INO1</i>	Lipid synthesis	Lipid	[2,24]

[&] All metabolites are located in the cytoplasm except those labeled with [m] (mitochondrial metabolites) or [e] (extracellular metabolites).

* The “and” logic implies protein complexes or multi-subunit proteins where each enzyme is required to catalyze a reaction; the minimum expression value of all enzymes is used to define the upper bound of reaction constraints. The “or” logic implies isozymes where each enzyme can individually catalyze a reaction; the maximum expression value of all enzymes is used to define the upper bound of reaction constraints.

^ Each objective function is defined by $C^T V$, where V is a vector of reaction fluxes and C is a vector of zeros with a value of one at the position of reactions of interest, as stated in the column. The exceptions are three ATP objectives: positive numbers of yielded ATPs are coefficients for reactions participating in ATP production, positive numbers of used ATPs are coefficient for reactions participating in ATP consumption, and positive numbers of yielded ATPs and negative numbers of used ATPs are entered as coefficients for the objective function of net ATP production. The ATP coefficients are in parentheses immediately following the name of the ATP-related objective functions. Other energy molecules are converted to the ATP unit: 1NADH=3ATP, 1NADPH=3ATP, 1FADH₂=2ATP, 1GTP=1ATP.

[§] This reaction represents three independent reactions catalyzed by three enzymes, thus the average expression value of the three enzymes is used to define the upper bound of reaction constraints [15].

[#] The flux of this reaction is set to 0 because *SFC1* encodes an antiporter, which transports cytoplasmic succinate into mitochondria in exchange for fumarate [19].

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