

1. Imipenem inhibition zone test
2. PCR validation of *mdtC* knockout strains

Column 15 is the wild-type strain control, and the remaining columns are knockout strain verifications. The strains used in sequencing and experiments are in column 3 (the 5000bp marker in the left column is not considered). The primers used were *mdtC*-2 listed in Table S6.

3. PCR validation of *mdtD* knockout strains

The 1st column is the verification of *mdtD* in the wild-type strain, the 2-3 columns are the verification of the *mdtD* gene deletion strain; the 5th column is the verification of the *mdtC*-1 primer in the wild-type strain, and the 6-8 columns are *mdtC*-1 Verification of primers in gene deletion strains; column 9 is the verification of *mdtC*-2 primers in wild-type strains, columns 10-12 are verification of *mdtC*-2 primers in gene deletion strains(the 5000bp marker in the left column is not considered).

4. PCR validation of *macB* knockout strains

The 1st column is the verification of *mdtC*-1 in the wild-type strain, the 2-3 columns are the verification of the *mdtC*-1 gene deletion strain; the 5th column is the verification of the *mdtC*-2 in the wild-type strain, and the 6-8 columns are verification of *mdtC*-2 in gene deletion strains; column 9 is the verification of *macB* primers in wild-type strains, columns 10-16 are verification of *mdtC*-2 primers in gene deletion strains(the 5000bp markers in the figure are not considered).

5. PCR validation of *mdtE* and *mdtF* knockout strains

We knocked out both the *mdtE* and *mdtF* genes in the operon where *mdtF* is located, although only the *mdtF* gene deletion strain was used in this study. The 1st column is the verification of *mdtE* in the knockout strain, the 2nd column is the verification of *mdtE* in the wild-type strain; the 3rd column is the verification of *mdtF* in the knockout strain, and the 4th column is the verification of *mdtF* in the wild-type strain.

6. GO classification of differentially expressed genes

7. GO classification of differentially expressed genes

8. GO classification of differentially expressed genes

9. GO classification of differentially expressed genes

10. COG enrichment of differentially expressed genes

11. COG enrichment of differentially expressed genes

12. Enrichment of KEGG pathway for differentially expressed genes

13. Carbon fixation pathways

14. Carbon metabolism

15. Sulfur metabolism

16. Sulfur relay system

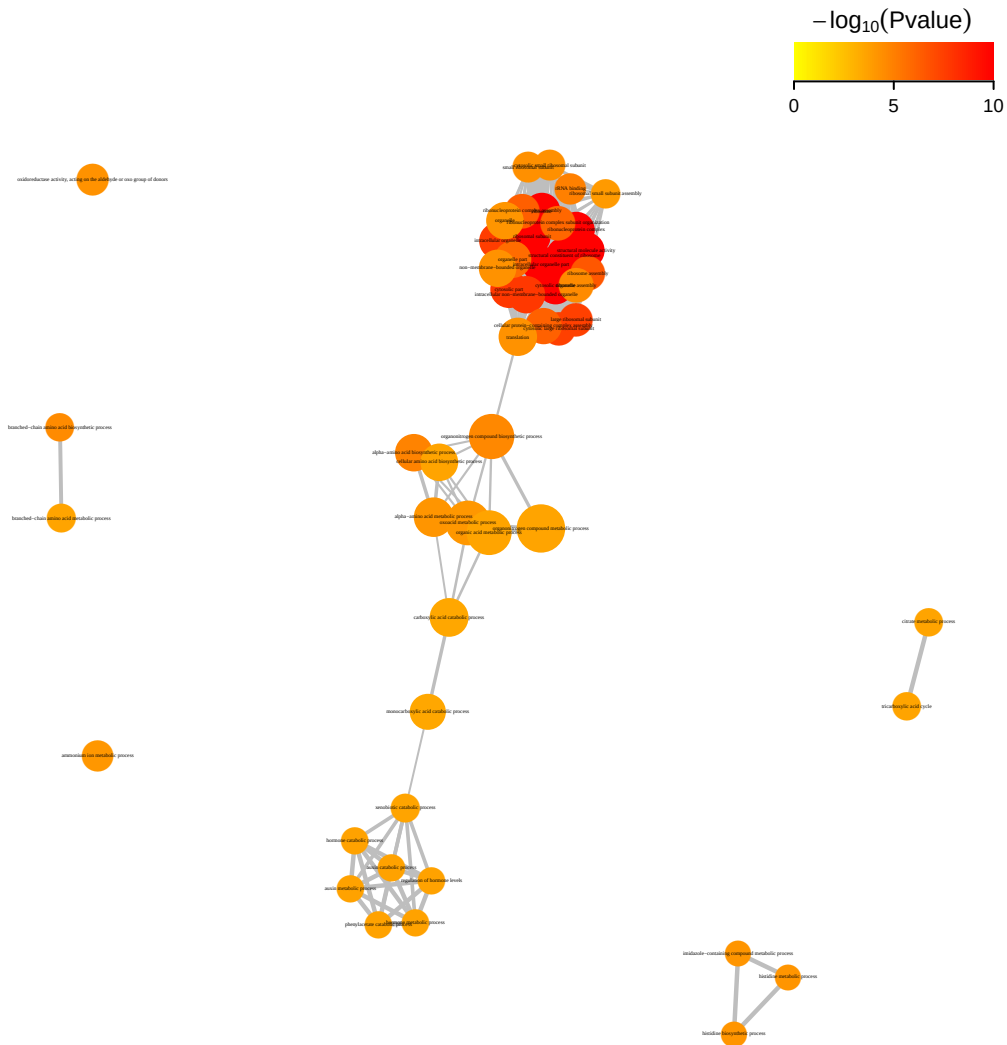
17. Nitrogen metabolism

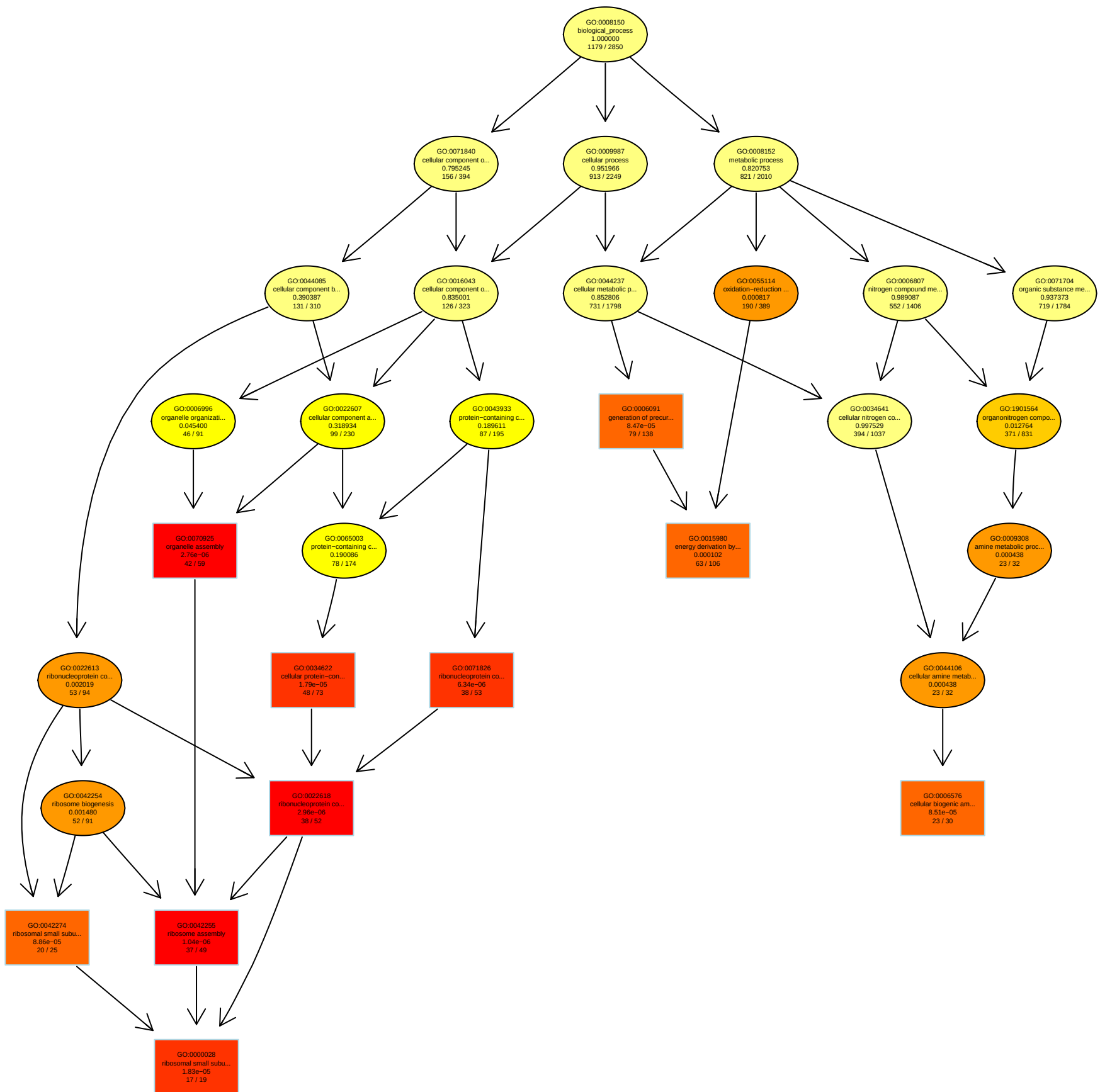
18. Starch and sucrose metabolism

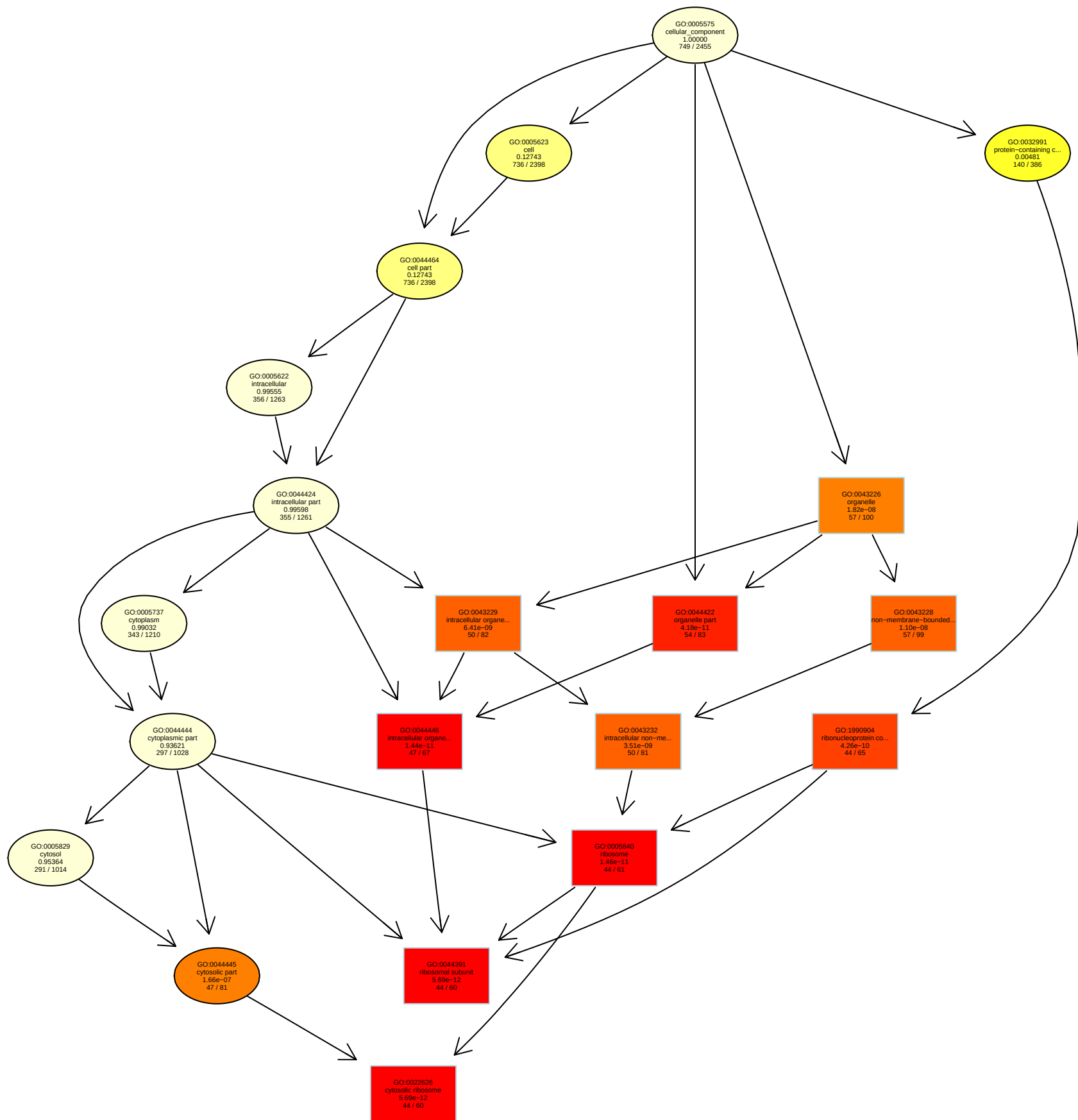
19. Glycolysis/Gluconeogenesis

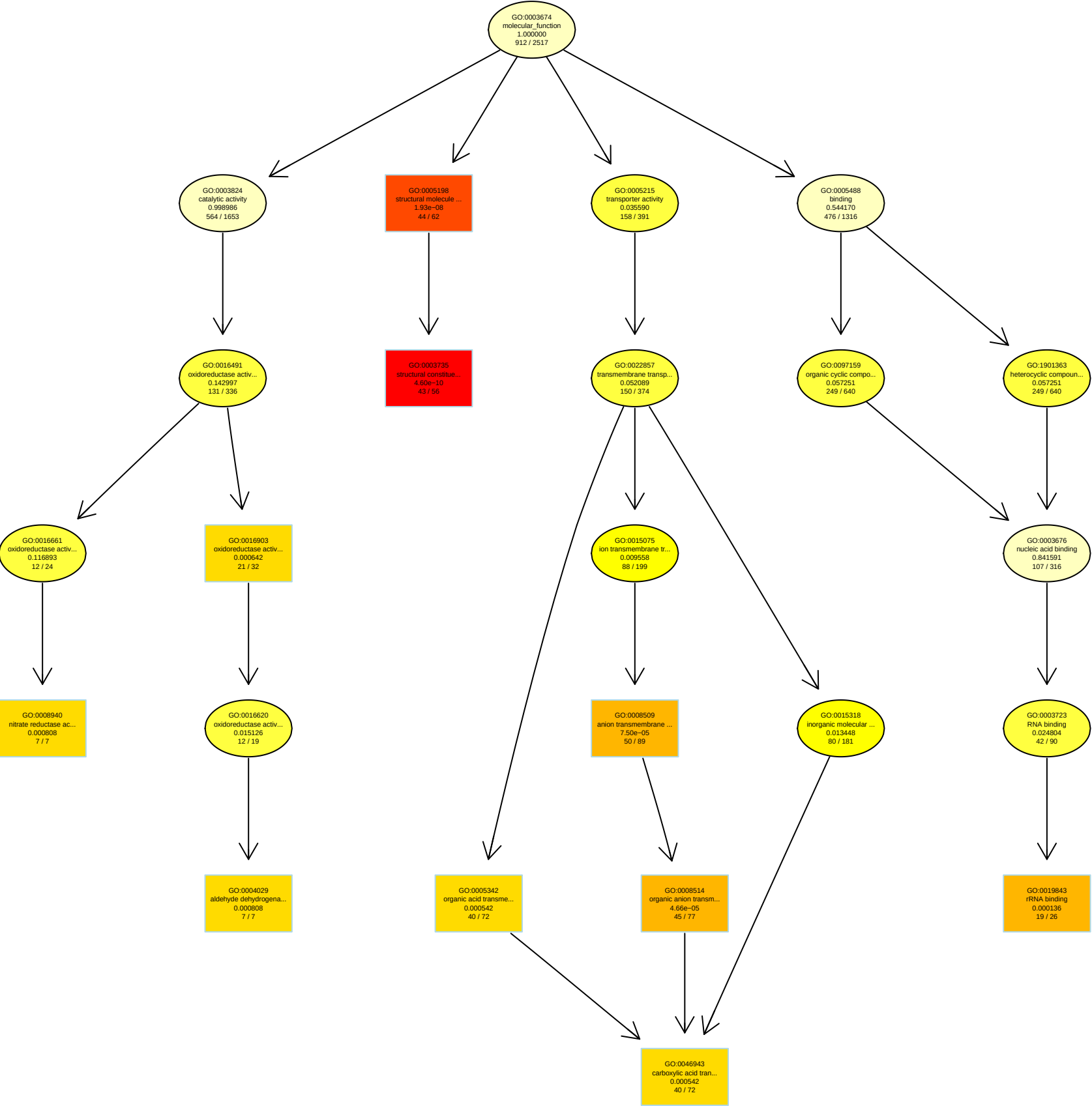
20. Pentose phosphate pathway

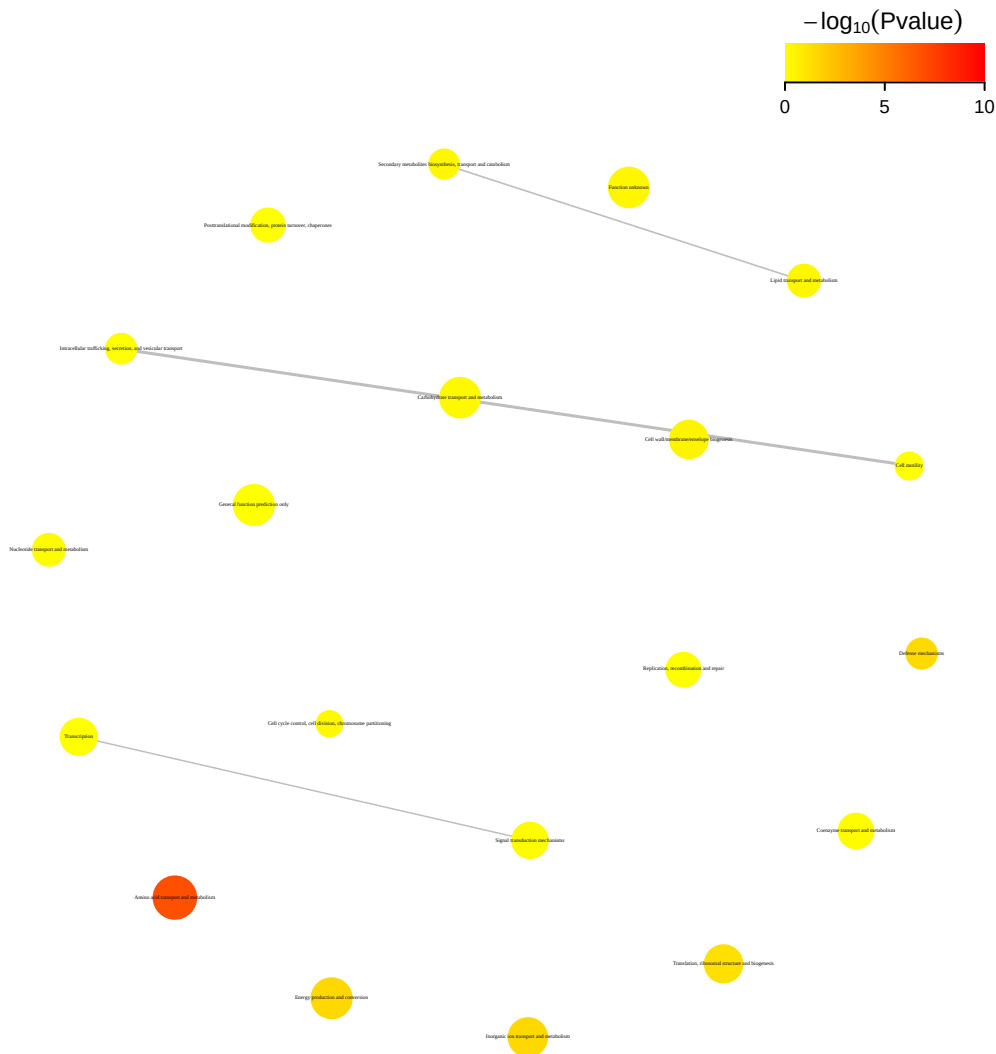
21. Phosphotransferase system (PTS)
22. Fatty acid degradation
23. Citrate cycle (TCA)
24. Biosynthesis of amino acid
25. Arginine and proline metabolism
26. Arginine biosynthesis
27. Lysine biosynthesis
28. Valine, leucine and isoleucine biosynthesis
29. Tryptophan metabolism
30. Ribosome
31. Purine metabolism
32. Lipopolysaccharide biosynthesis
33. Peptidoglycan biosynthesis
34. ABC transporters
35. Phosphotransferase system (PTS)
36. Two-component system
37. Flagellar assembly
38. Quorum sensing
39. Biofilm formation
40. Bacterial chemotaxis

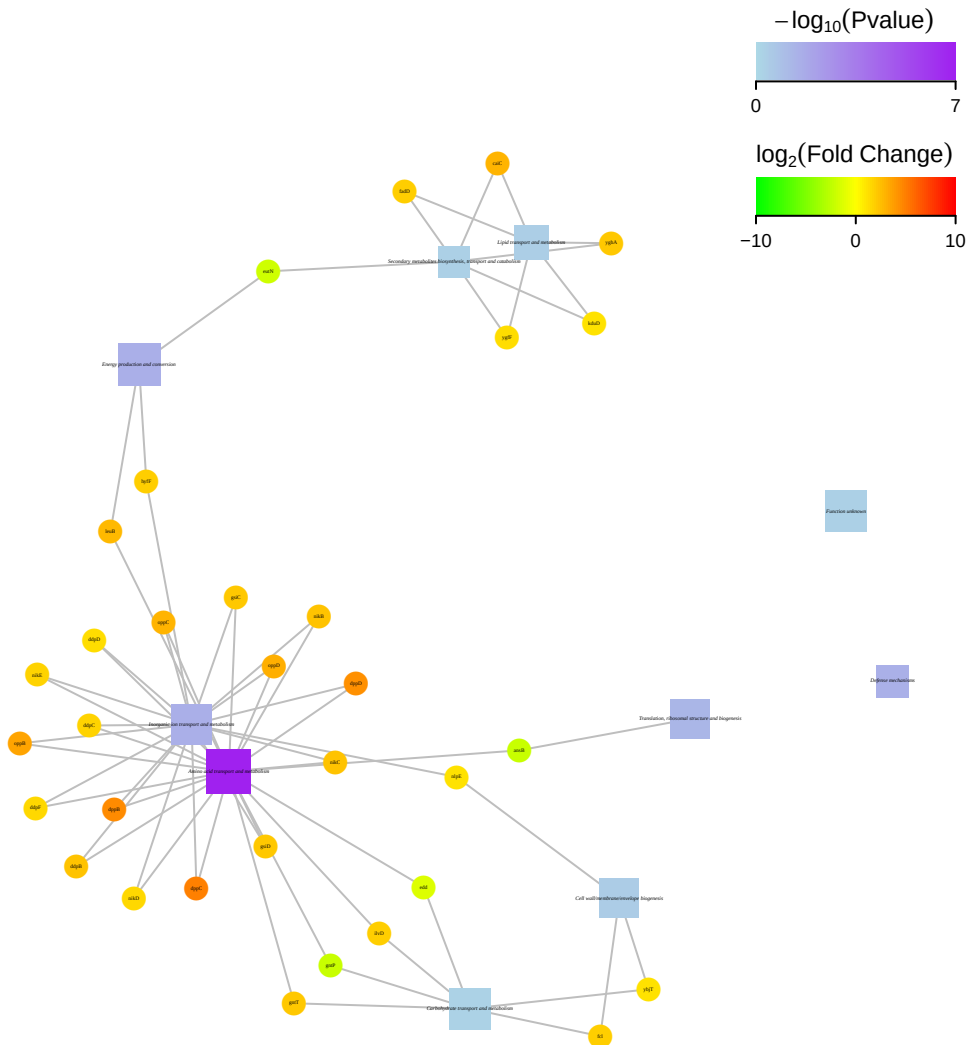


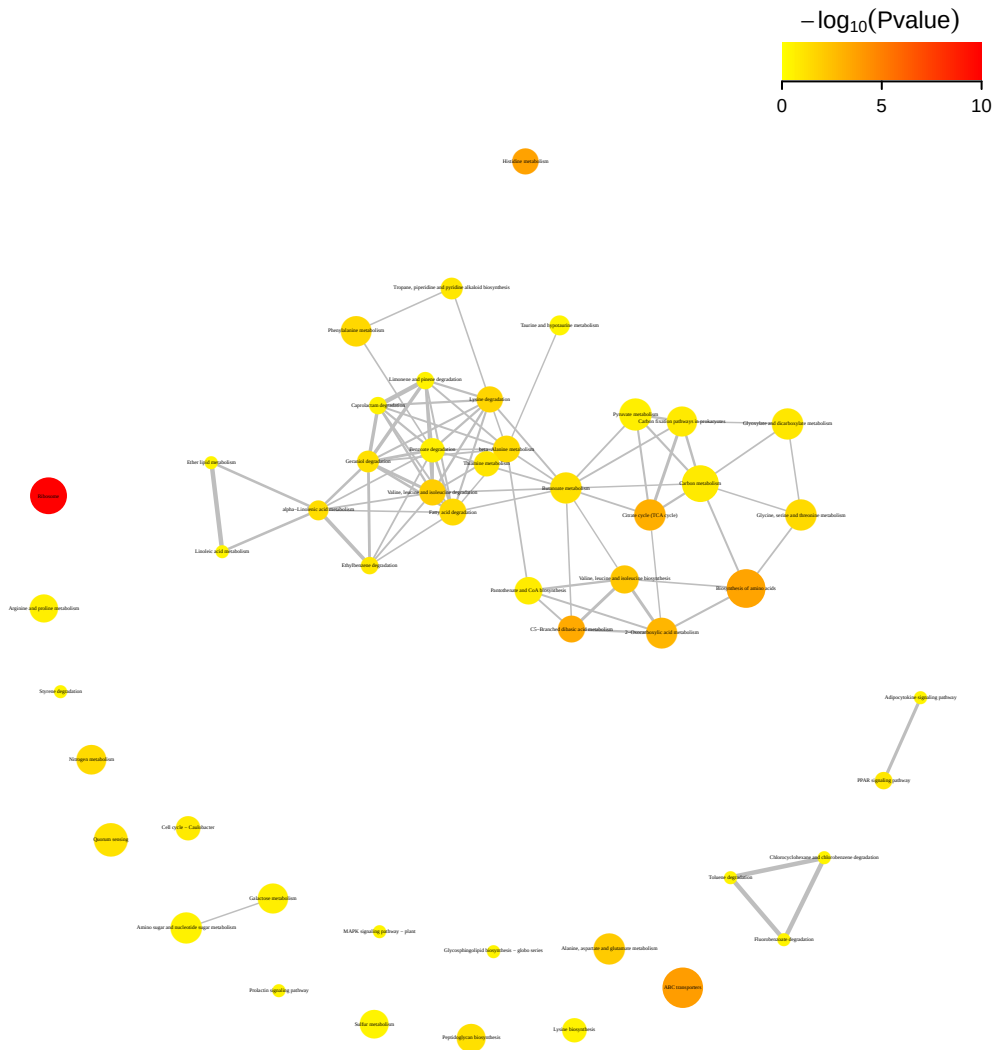




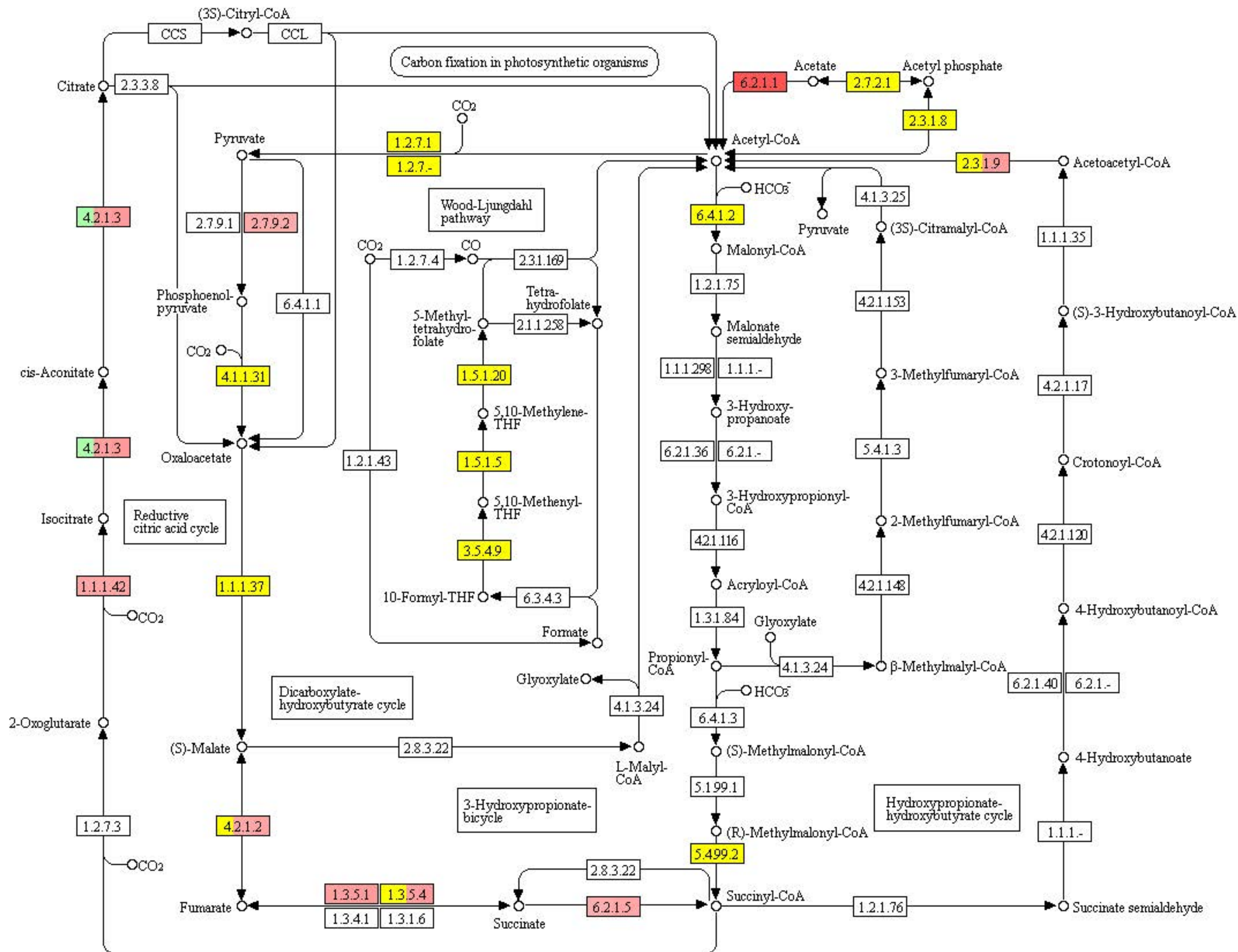


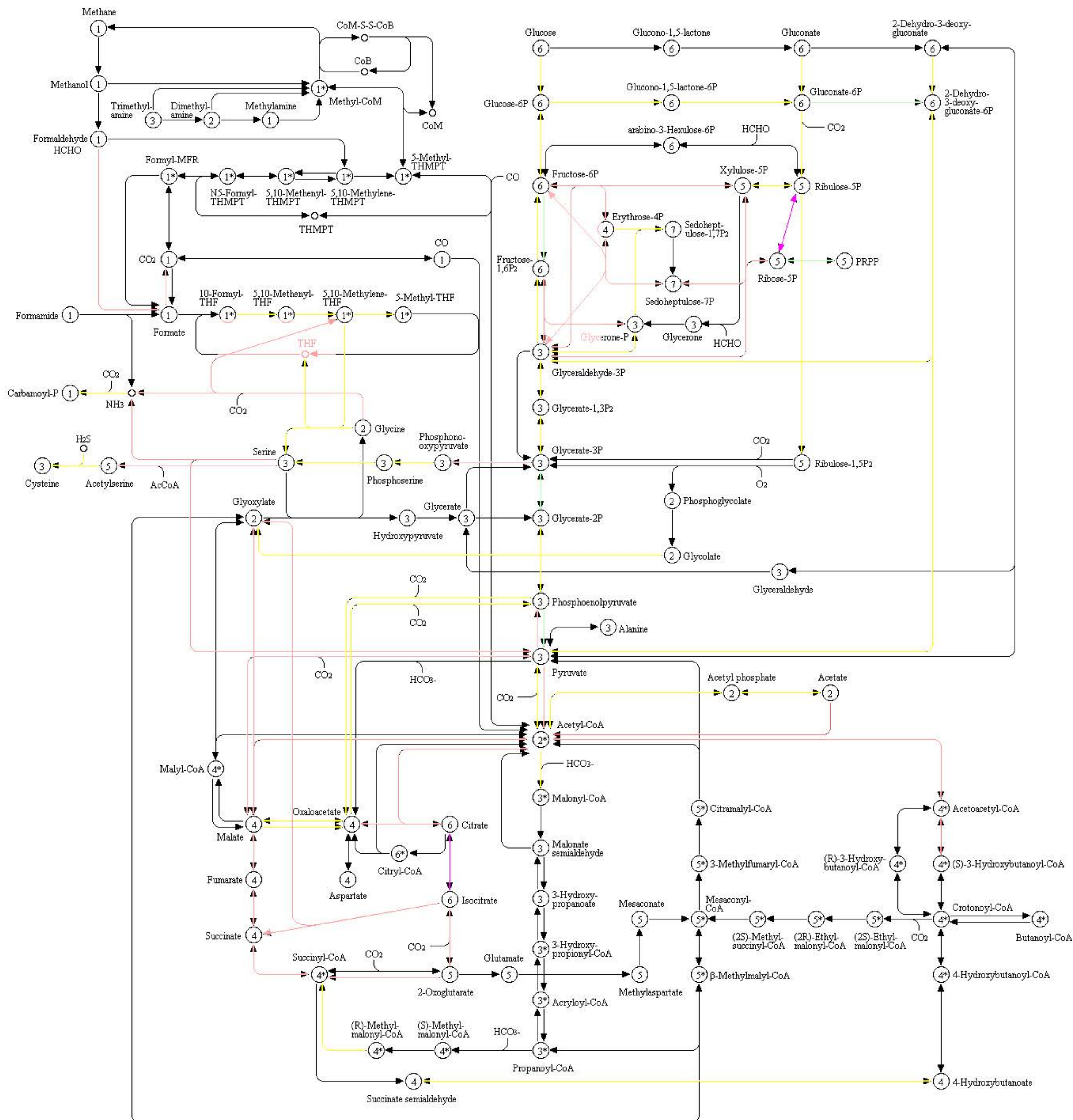




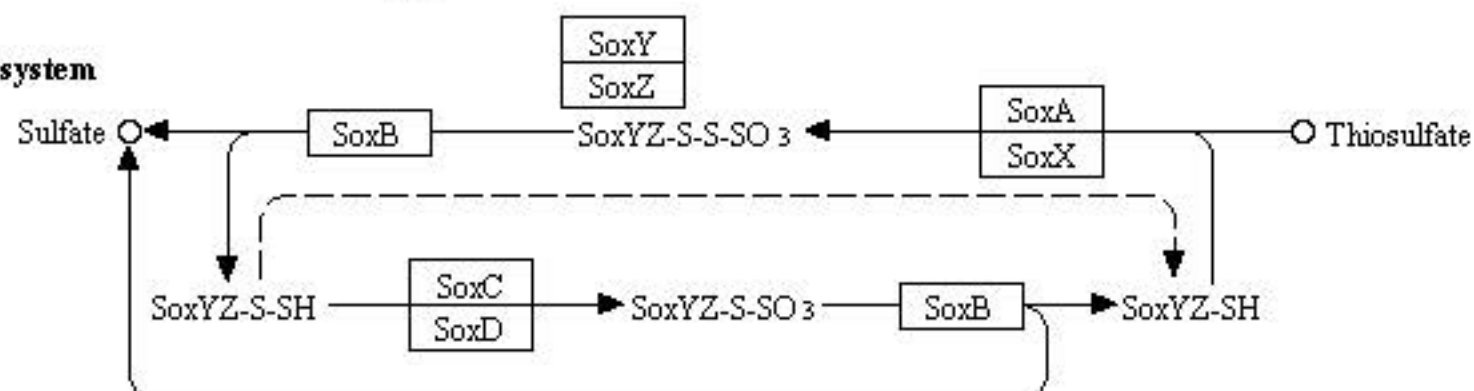
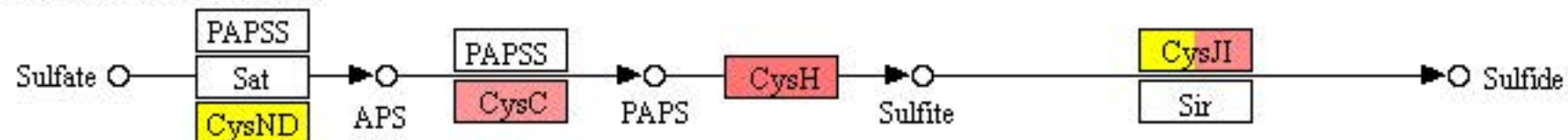


CARBON FIXATION PATHWAYS IN PROKARYOTES

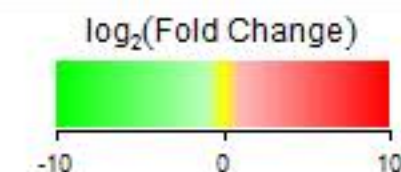




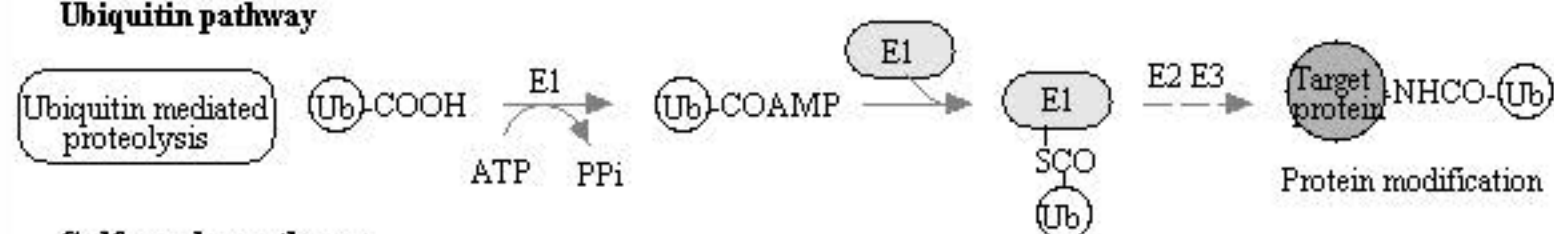
Oxidation state +6



SULFUR RELAY SYSTEM

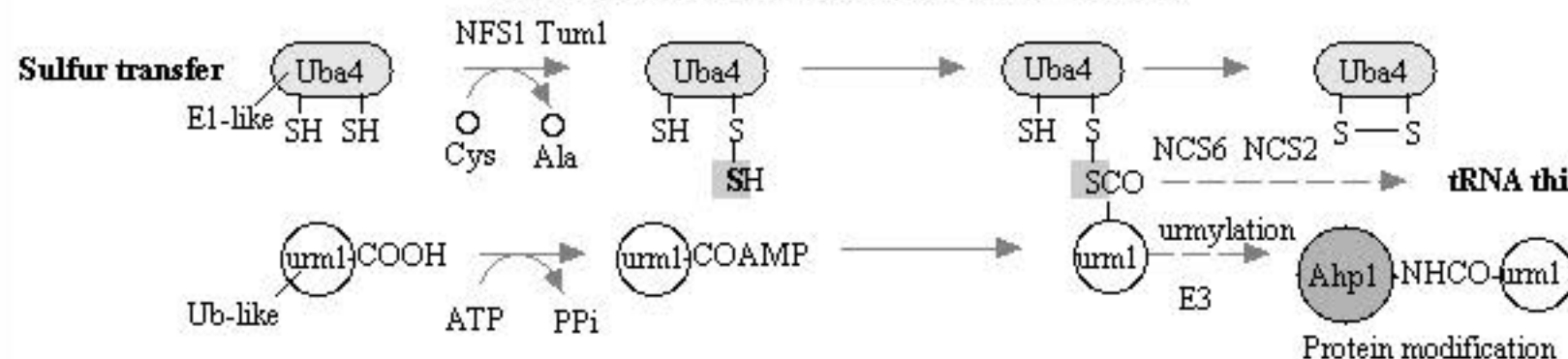


Ubiquitin pathway

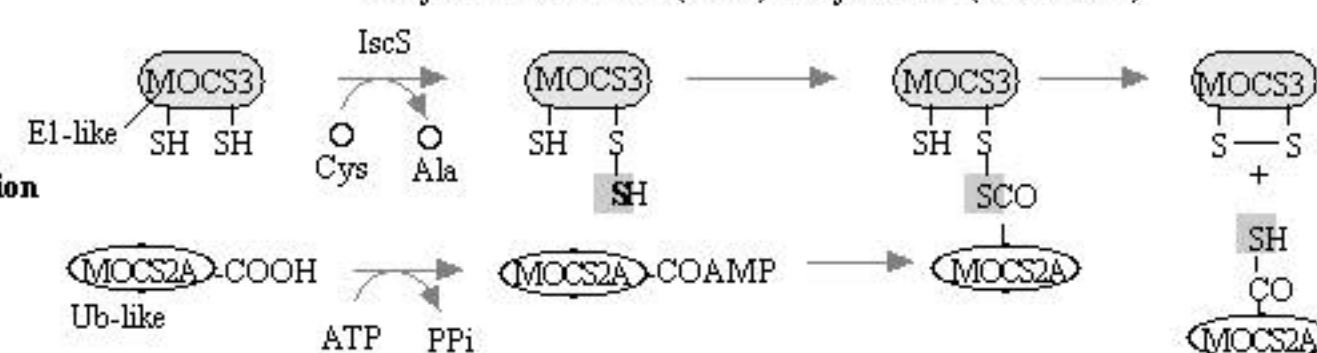


Sulfur-relay pathway

2-thiouridine biosynthesis (*Saccharomyces cerevisiae*)

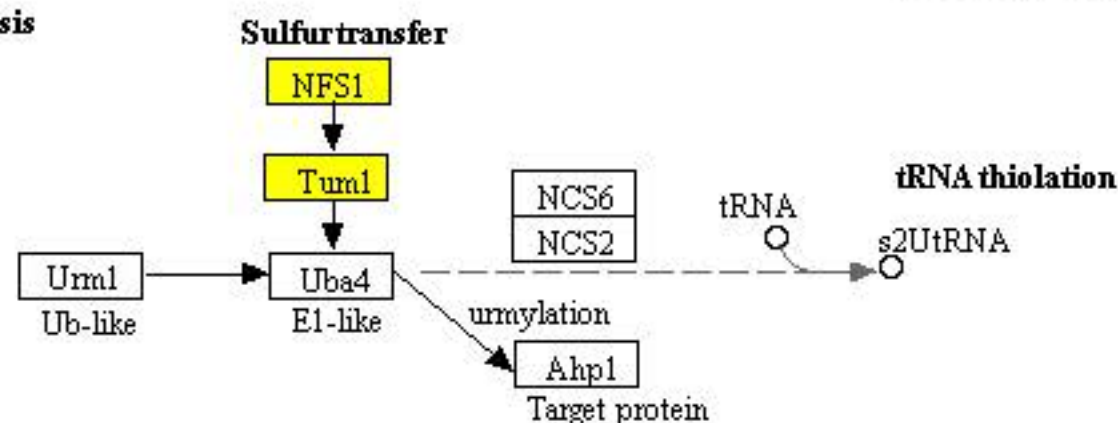


Molybdenum cofactor (Moco) biosynthesis (Mammals)

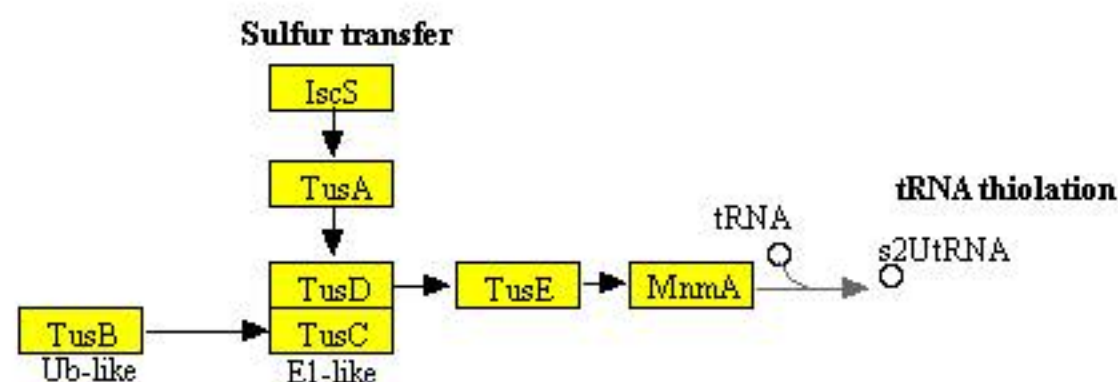


2-thiouridine biosynthesis

Eukaryote

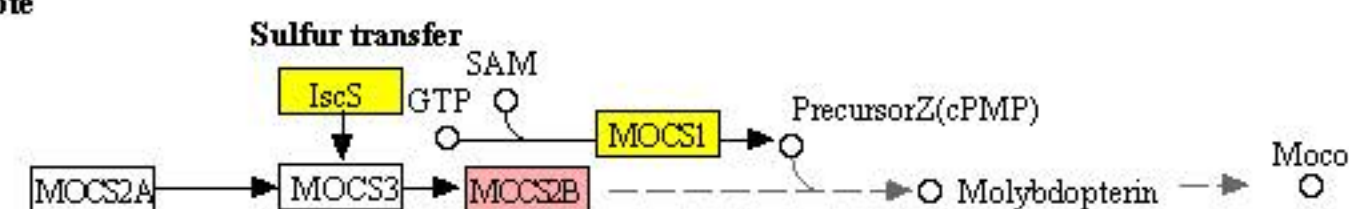


Prokaryote

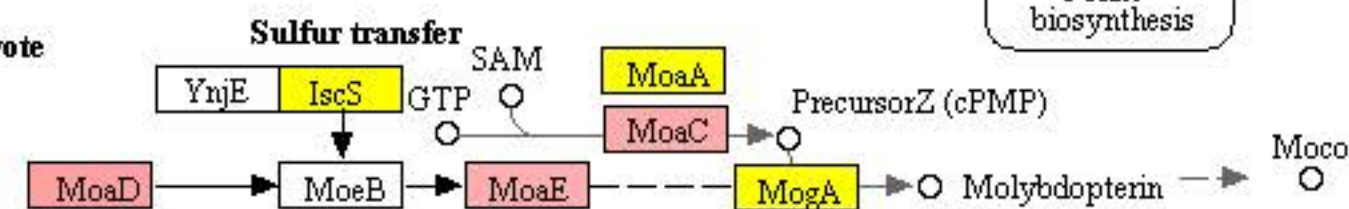


Moco biosynthesis

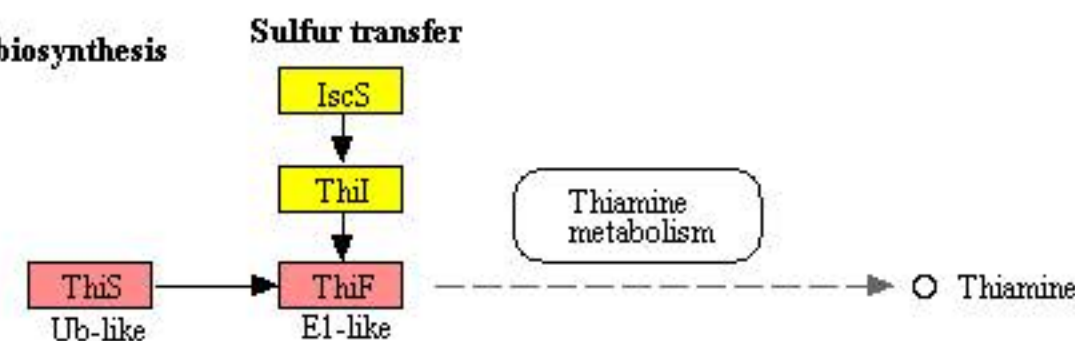
Eukaryote



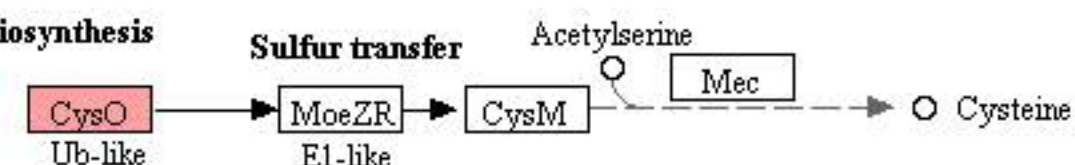
Prokaryote



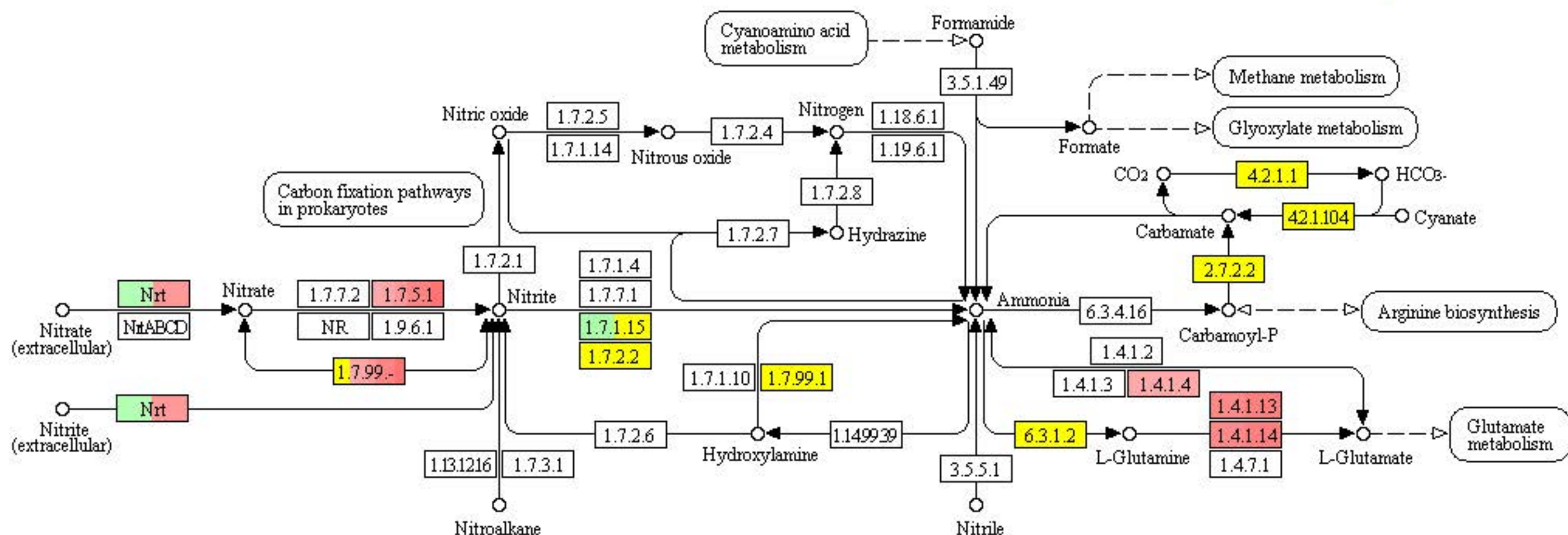
Thiamine biosynthesis



Cysteine biosynthesis

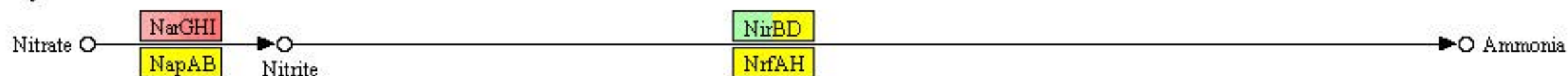


NITROGEN METABOLISM

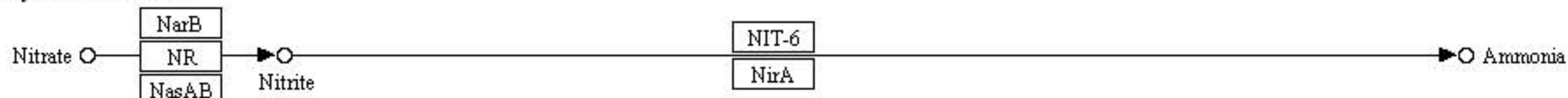


Oxidation state +5 +3 +2 +1 0 -1 -2 -3

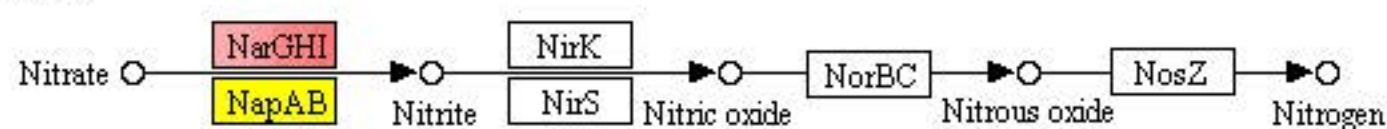
Dissimilatory nitrate reduction



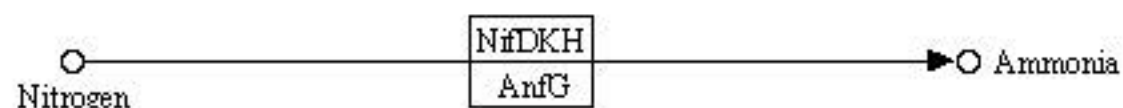
Assimilatory nitrate reduction



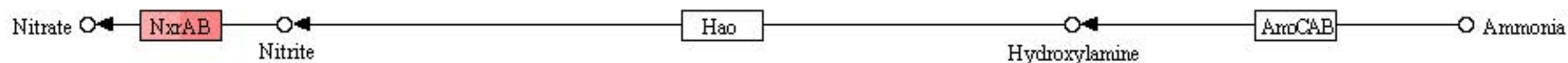
Denitrification



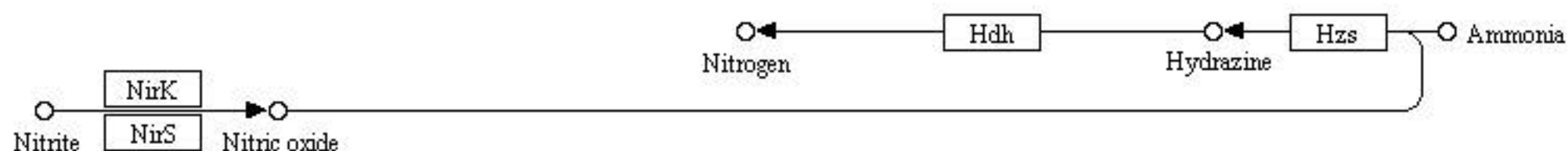
Nitrogen fixation



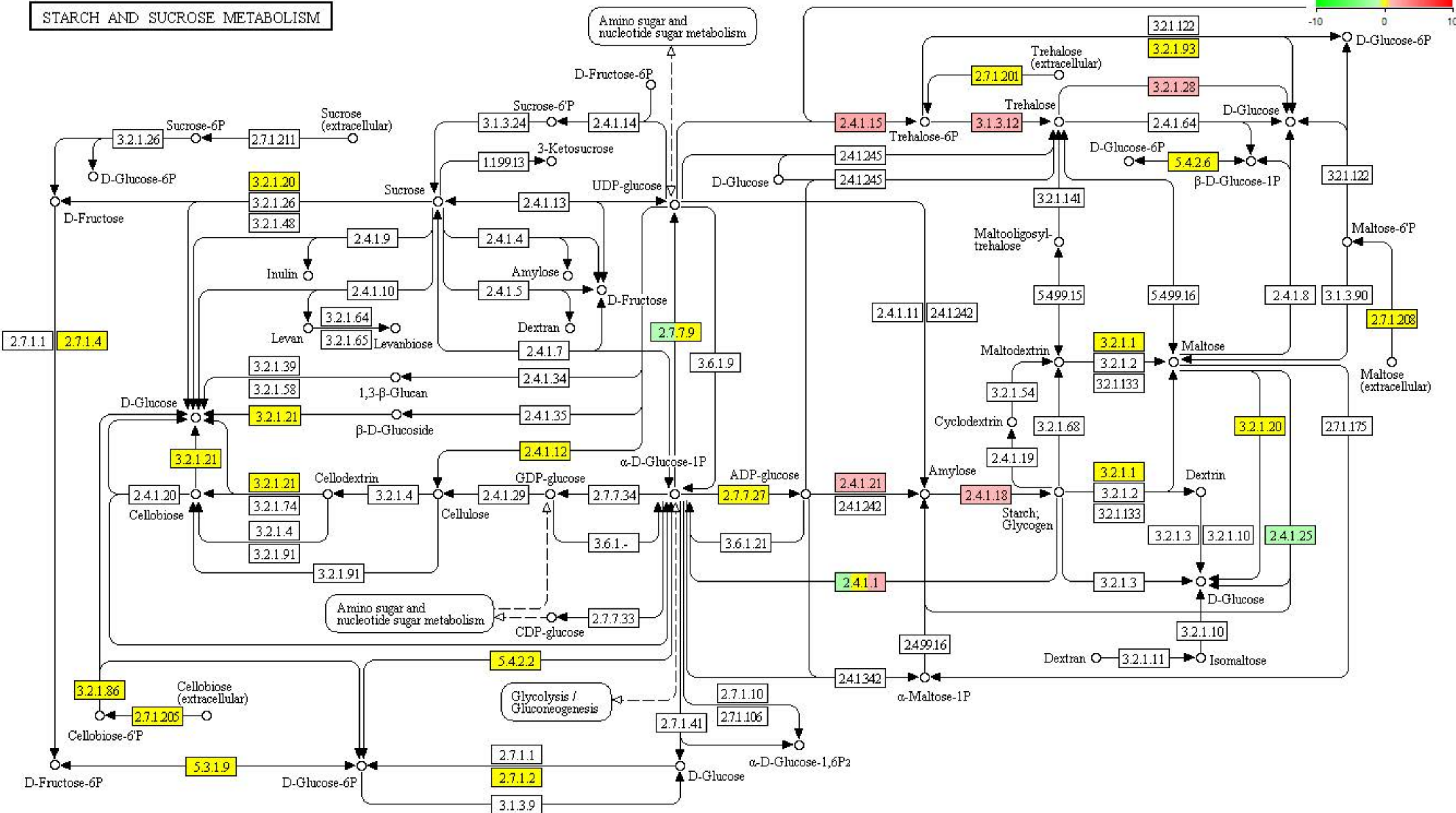
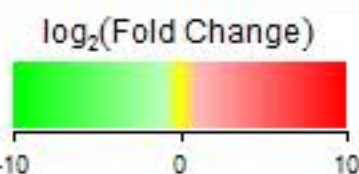
Nitrification



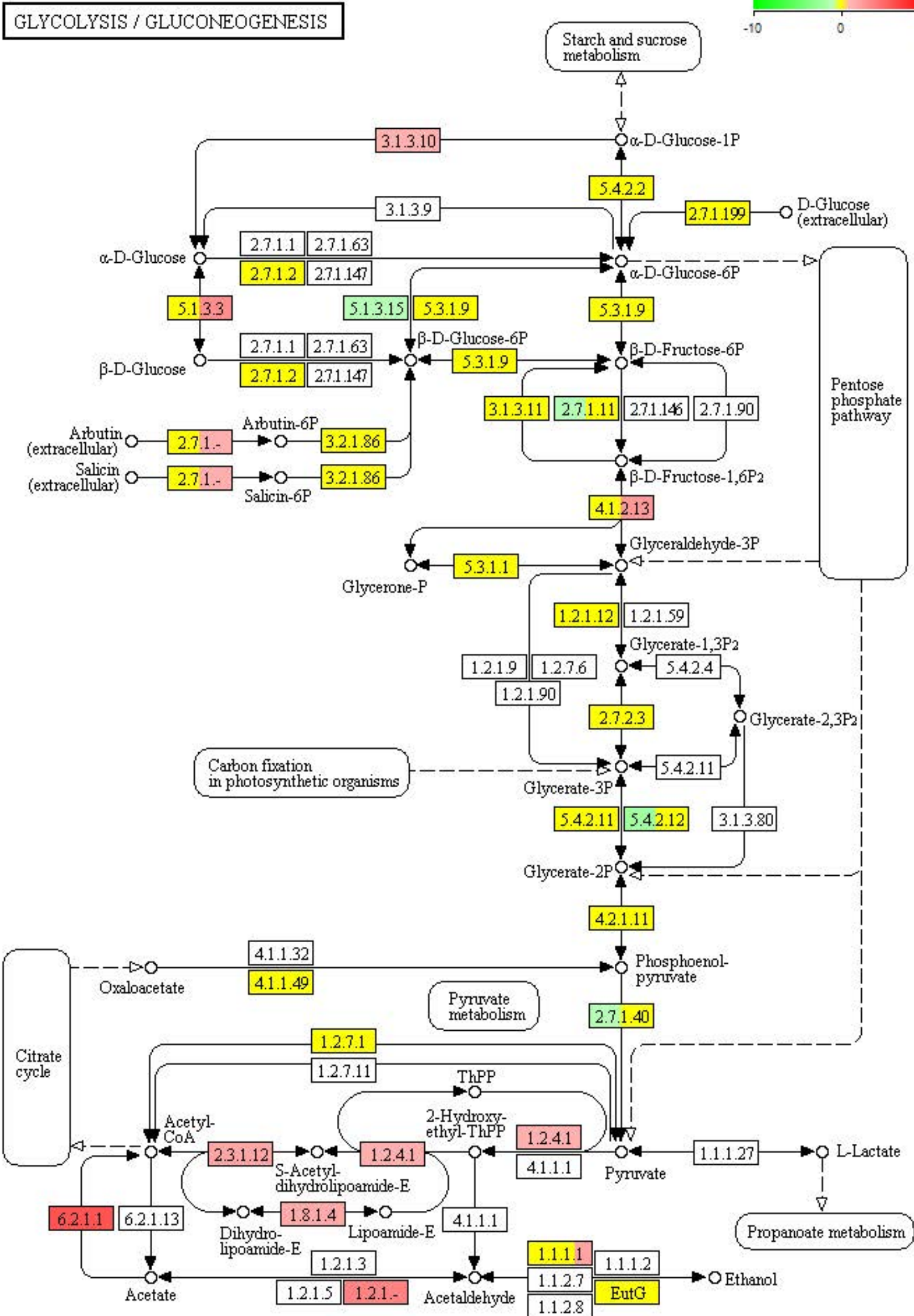
Anammox



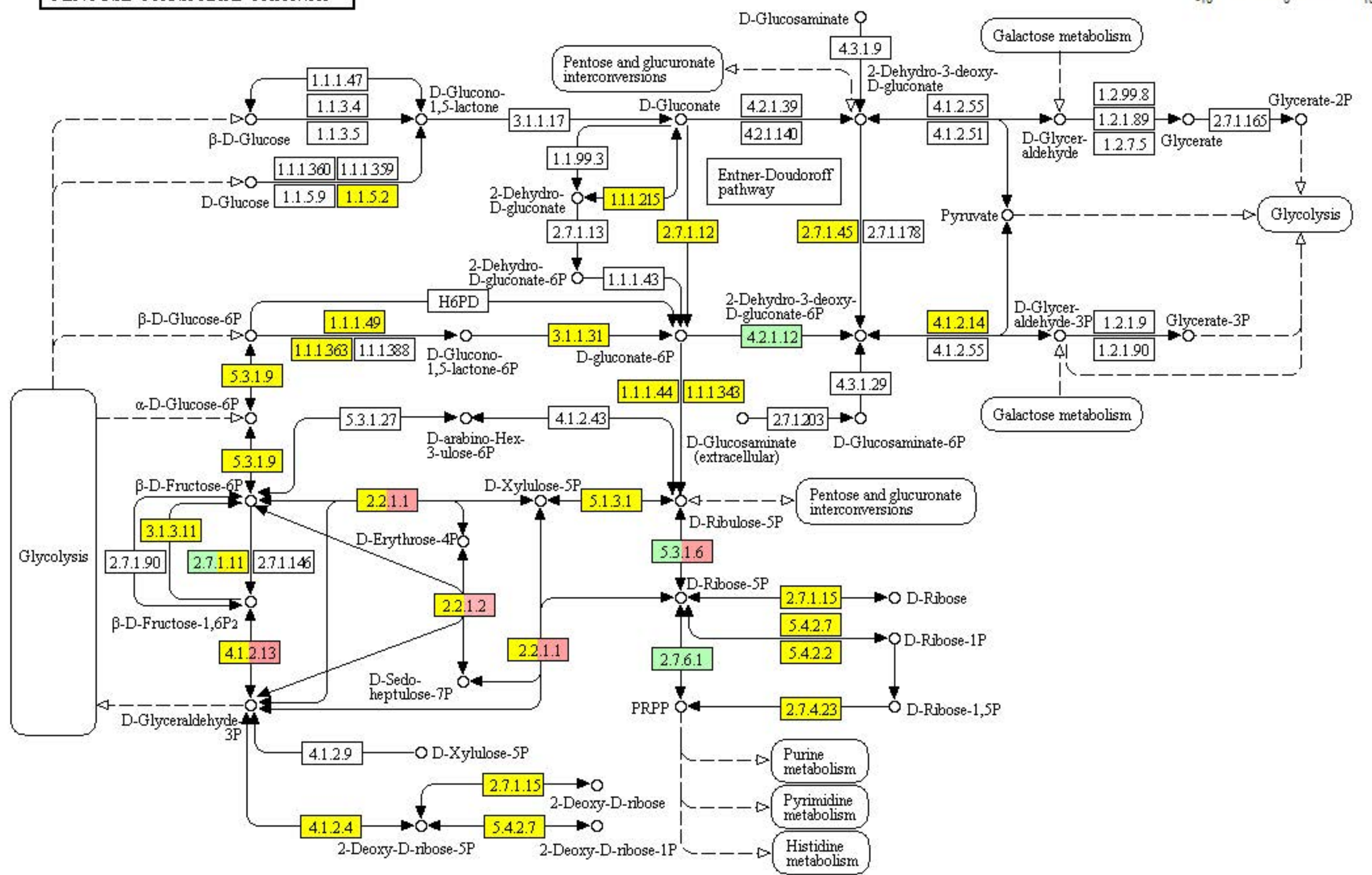
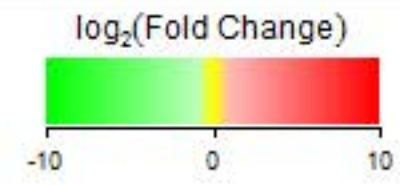
STARCH AND SUCROSE METABOLISM



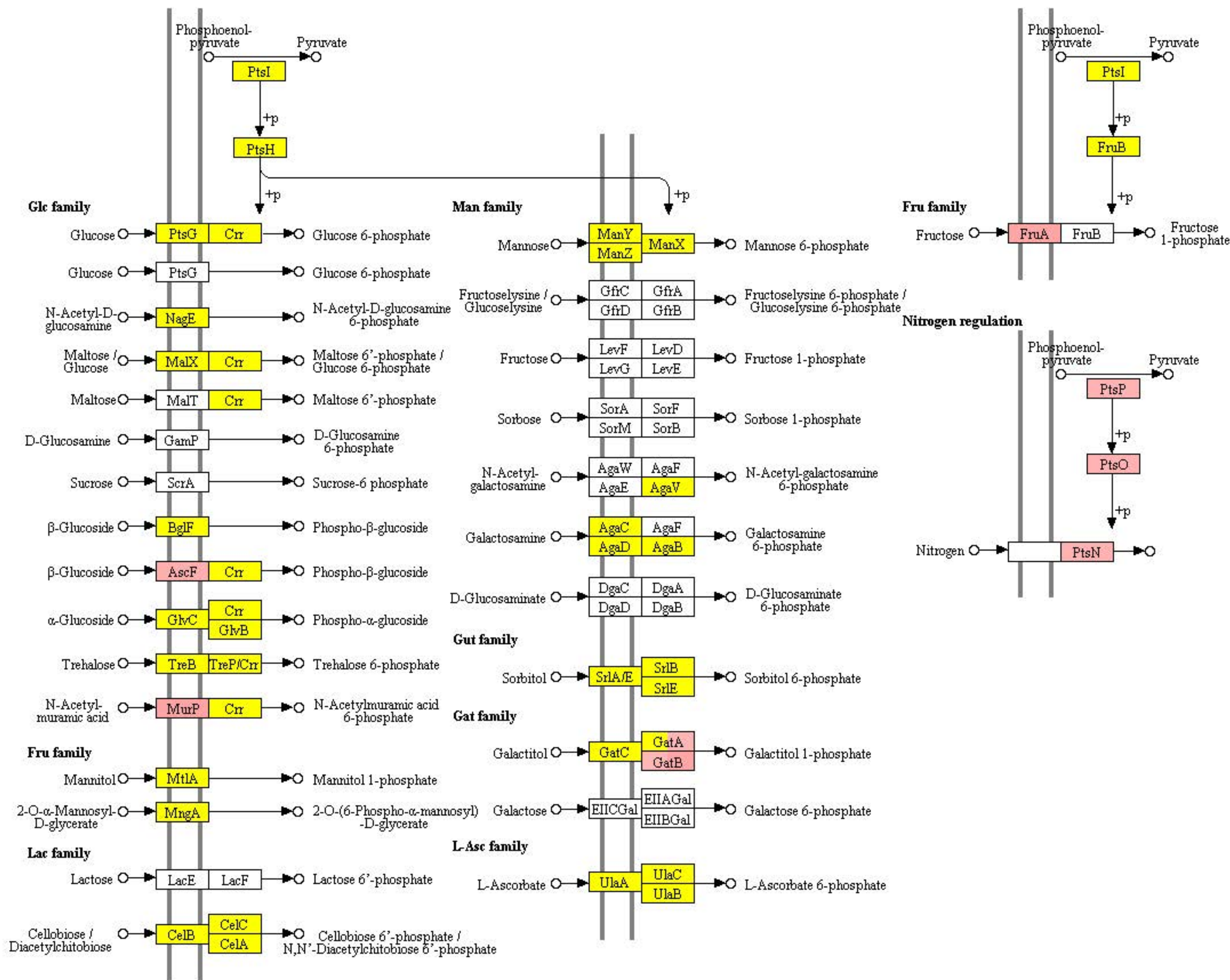
GLYCOLYSIS / GLUCONEOGENESIS



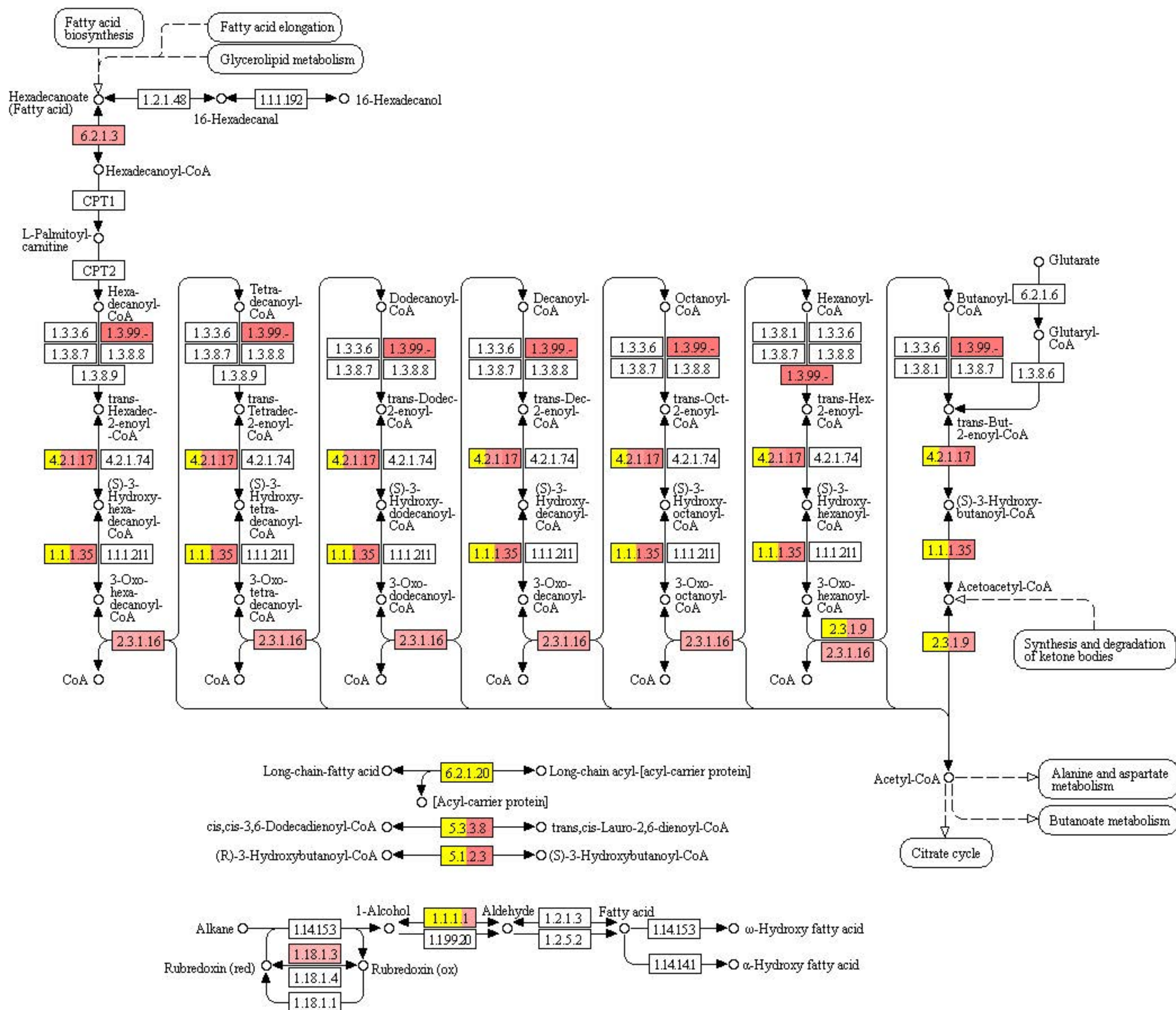
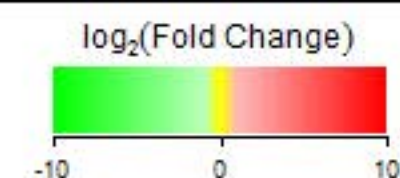
PENTOSE PHOSPHATE PATHWAY



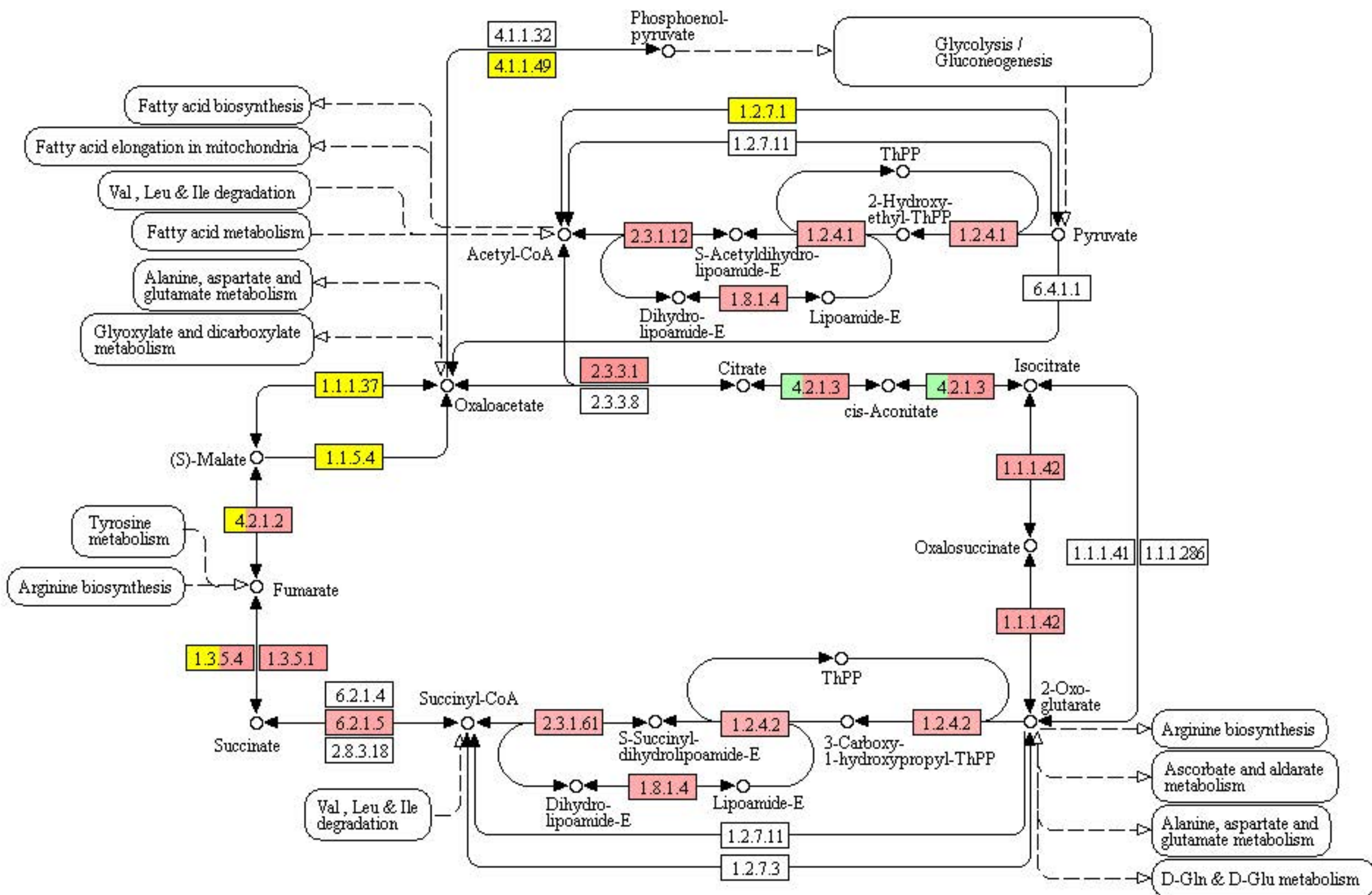
PHOSPHOTRANSFERASE SYSTEM (PTS)



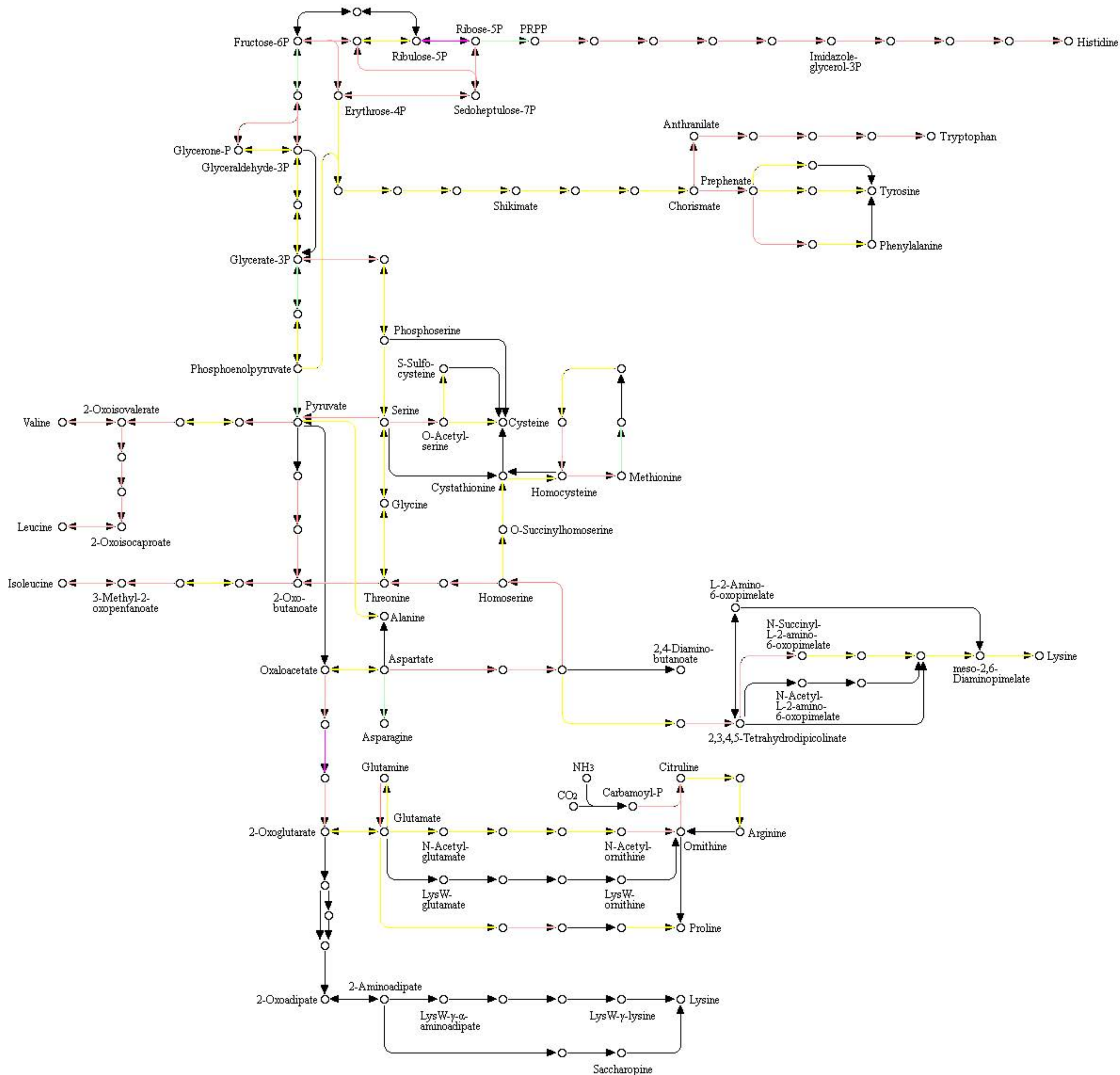
FATTY ACID DEGRADATION



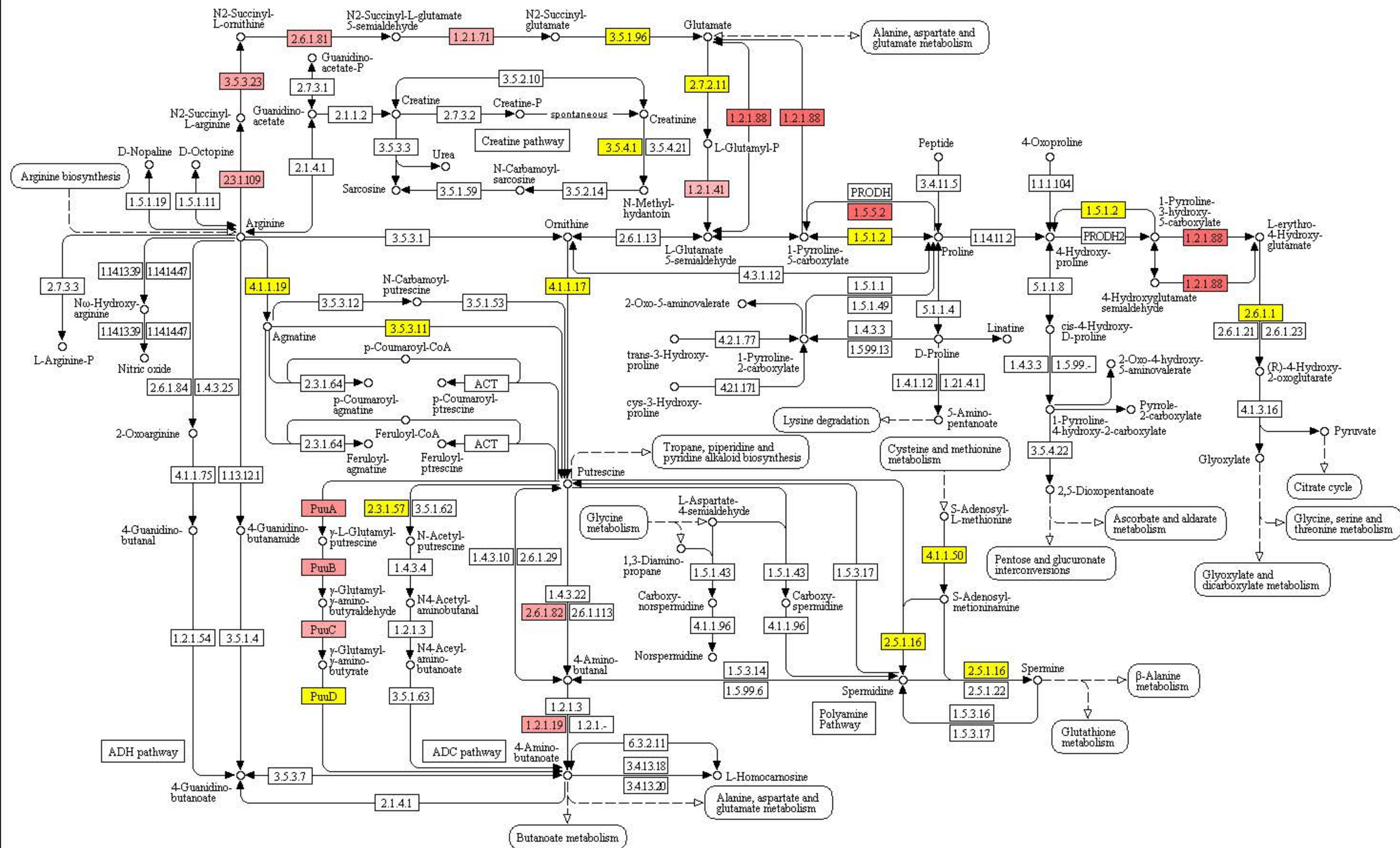
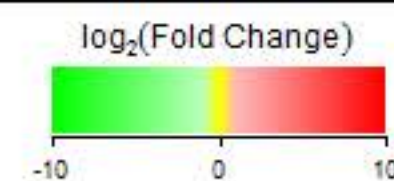
CITRATE CYCLE (TCA CYCLE)



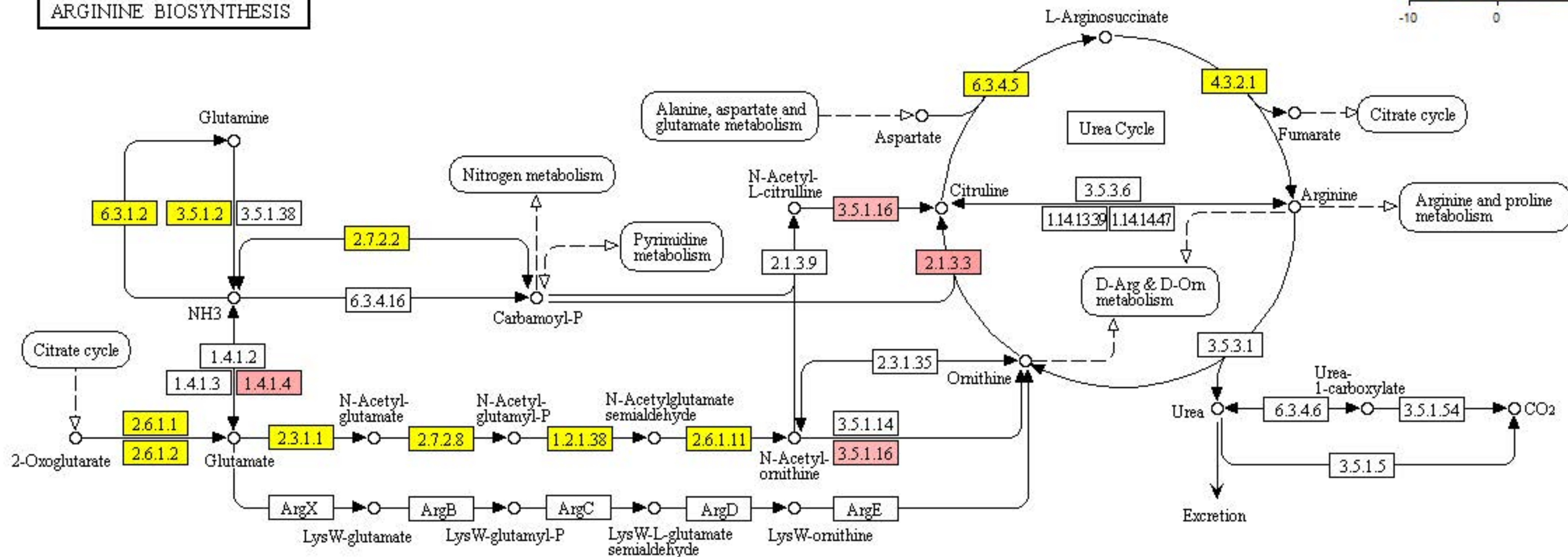
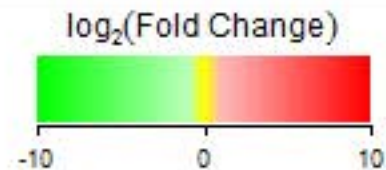
BIOSYNTHESIS OF AMINO ACIDS



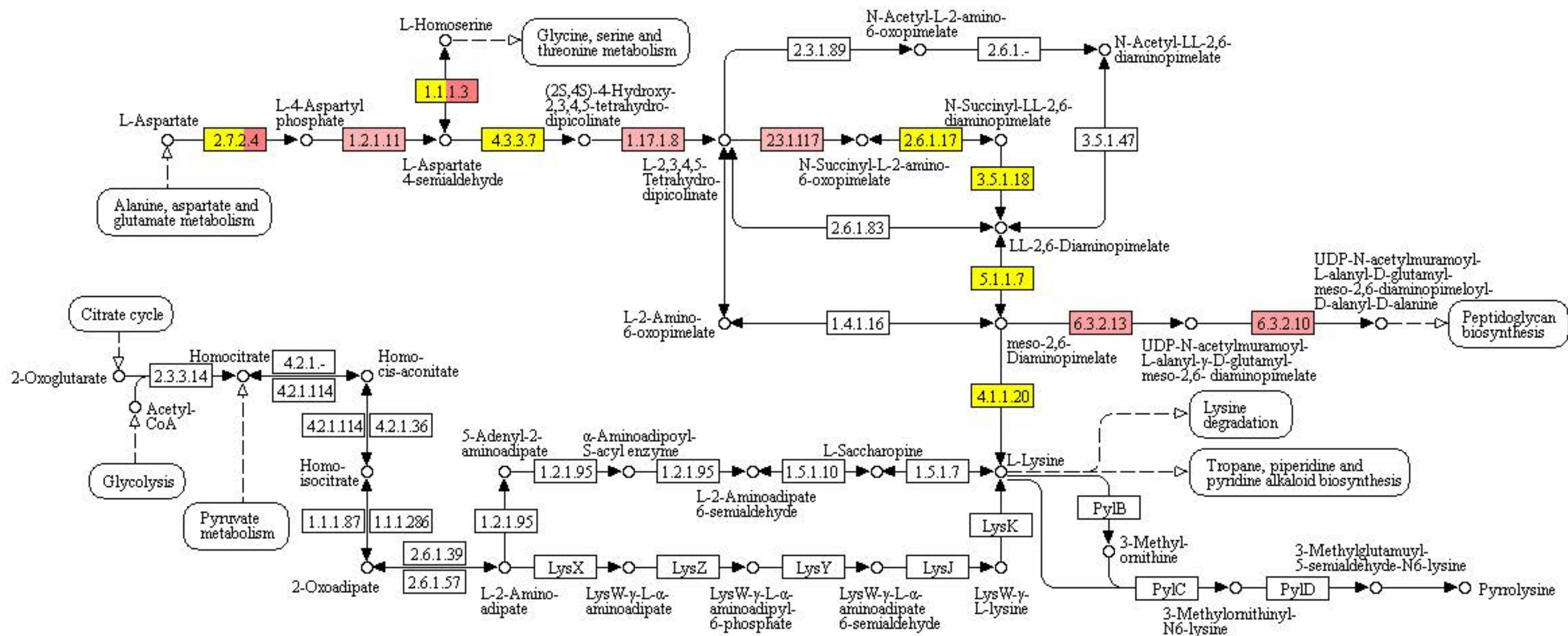
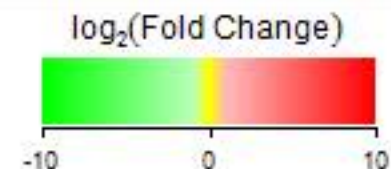
ARGININE AND PROLINE METABOLISM

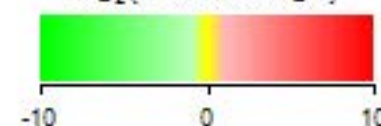


ARGININE BIOSYNTHESIS

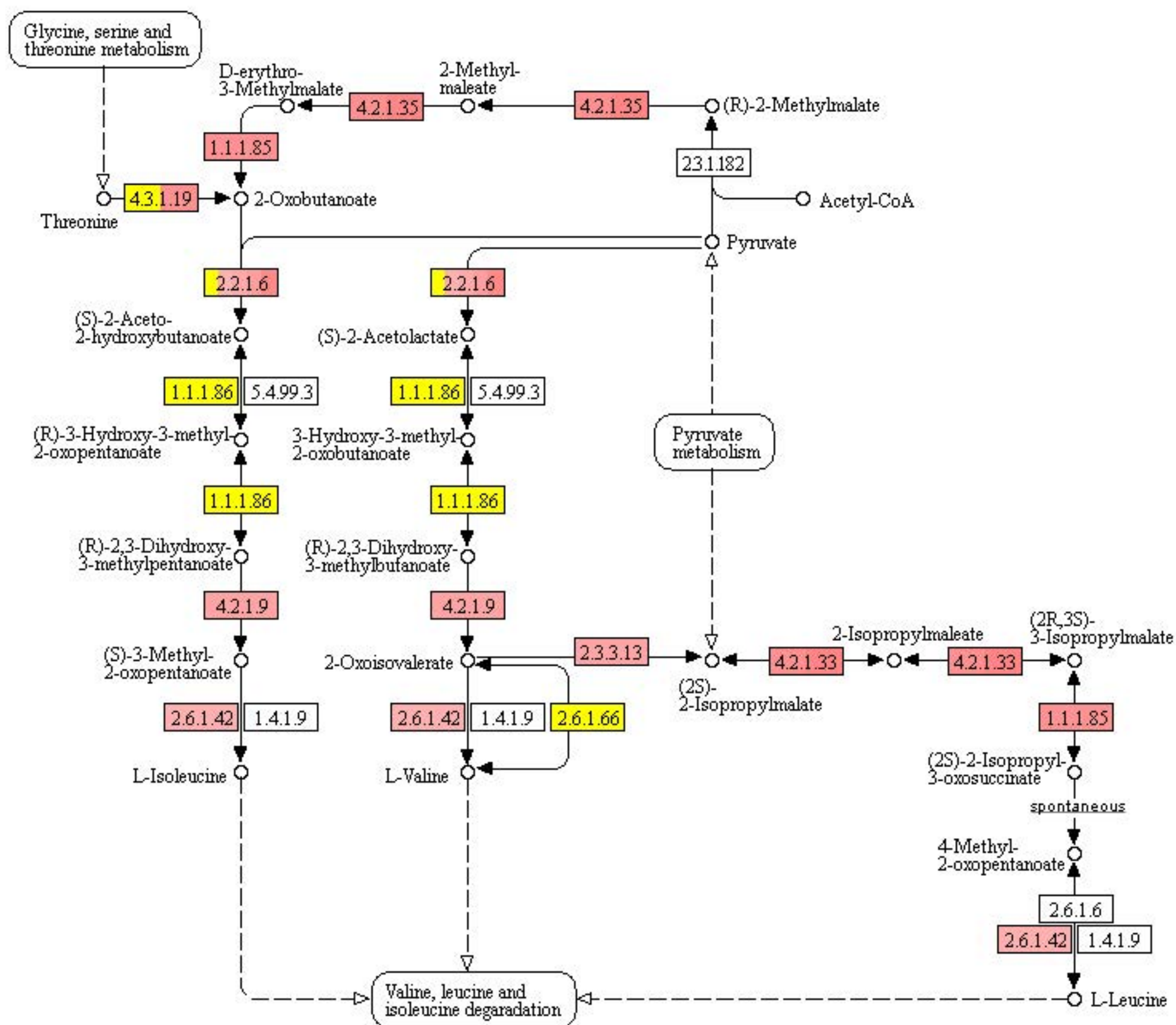


LYSINE BIOSYNTHESIS

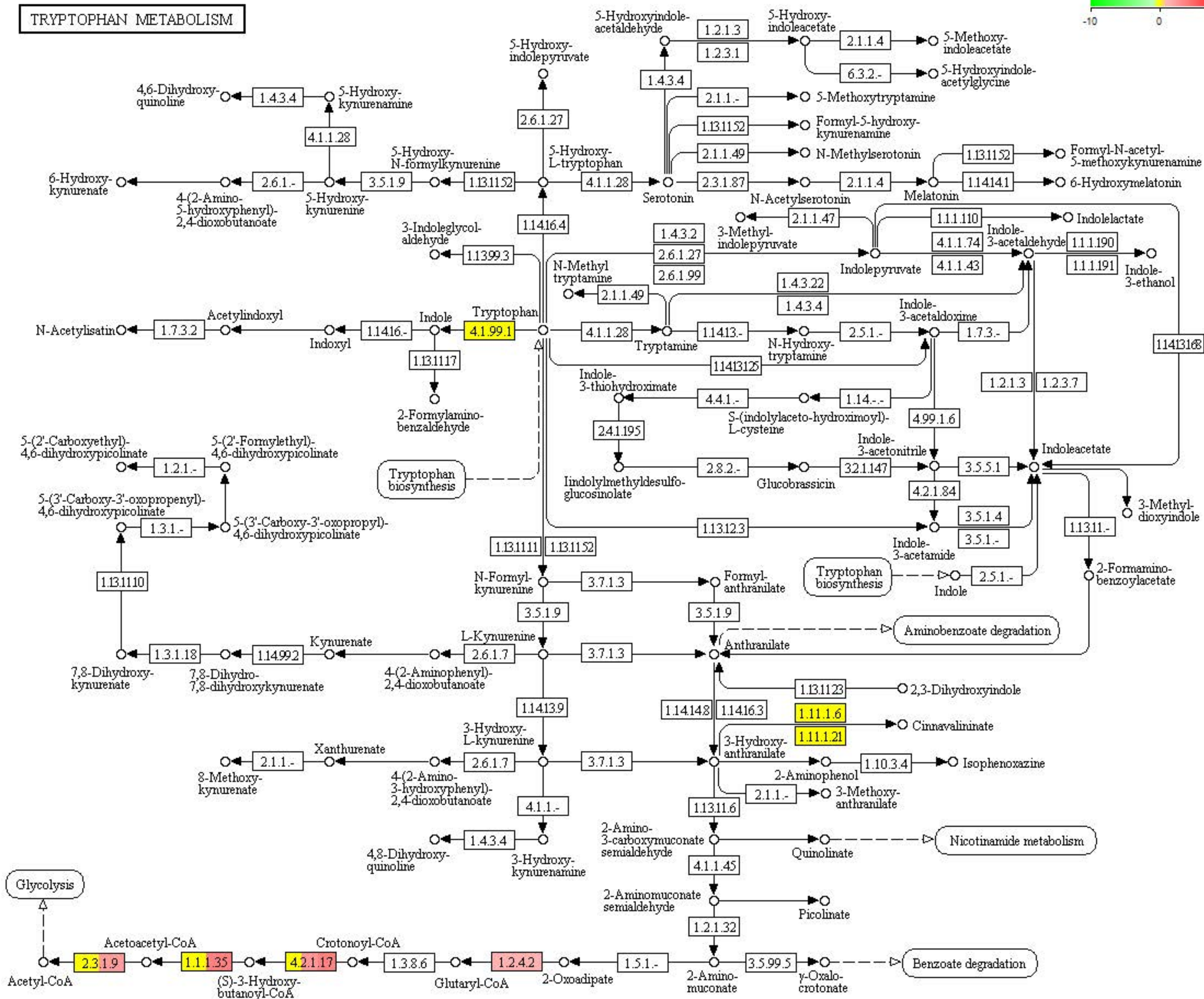
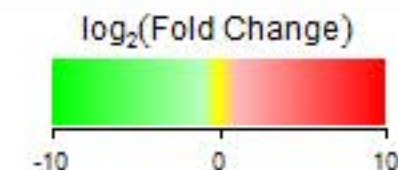




VALINE, LEUCINE AND ISOLEUCINE BIOSYNTHESIS

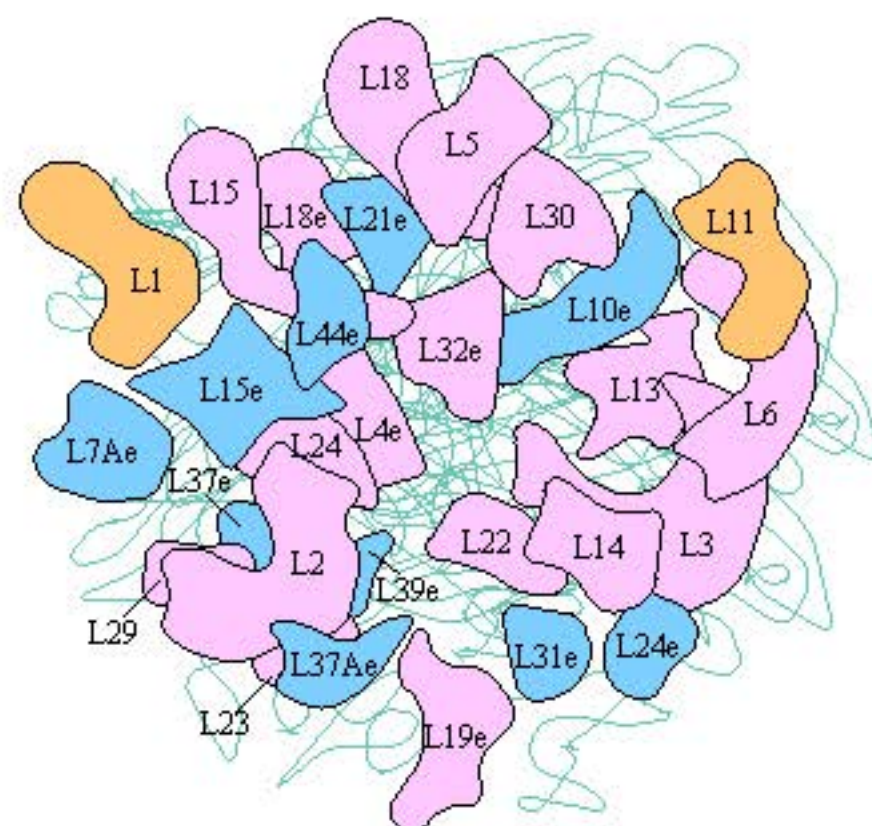


TRYPTOPHAN METABOLISM

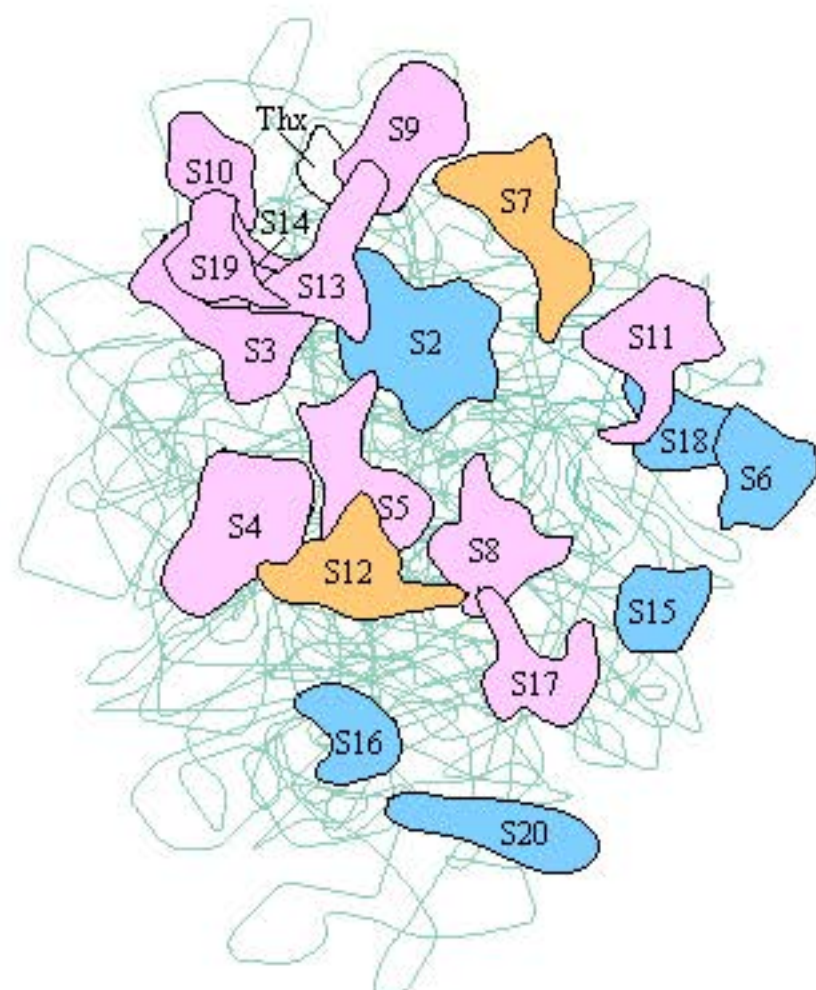




RIBOSOME



Large subunit (*Haloarcula marismortui*)



Small subunit (*Thermus aquaticus*)

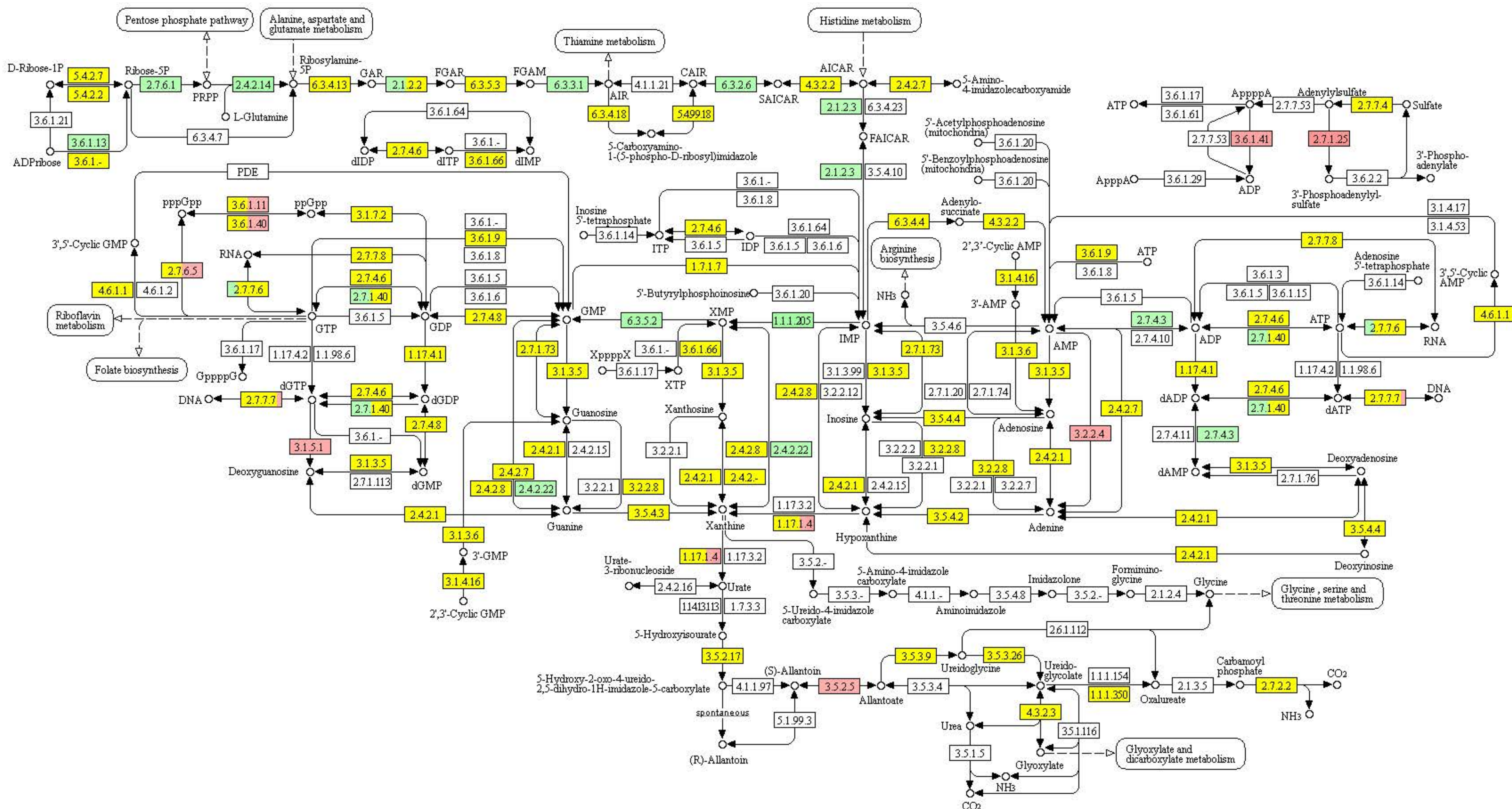
Ribosomal RNAs

Bacteria / Archaea Eukaryotes	23S	5S		16S
	25S	5S	5.8S	18S

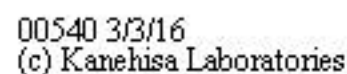
Ribosomal proteins

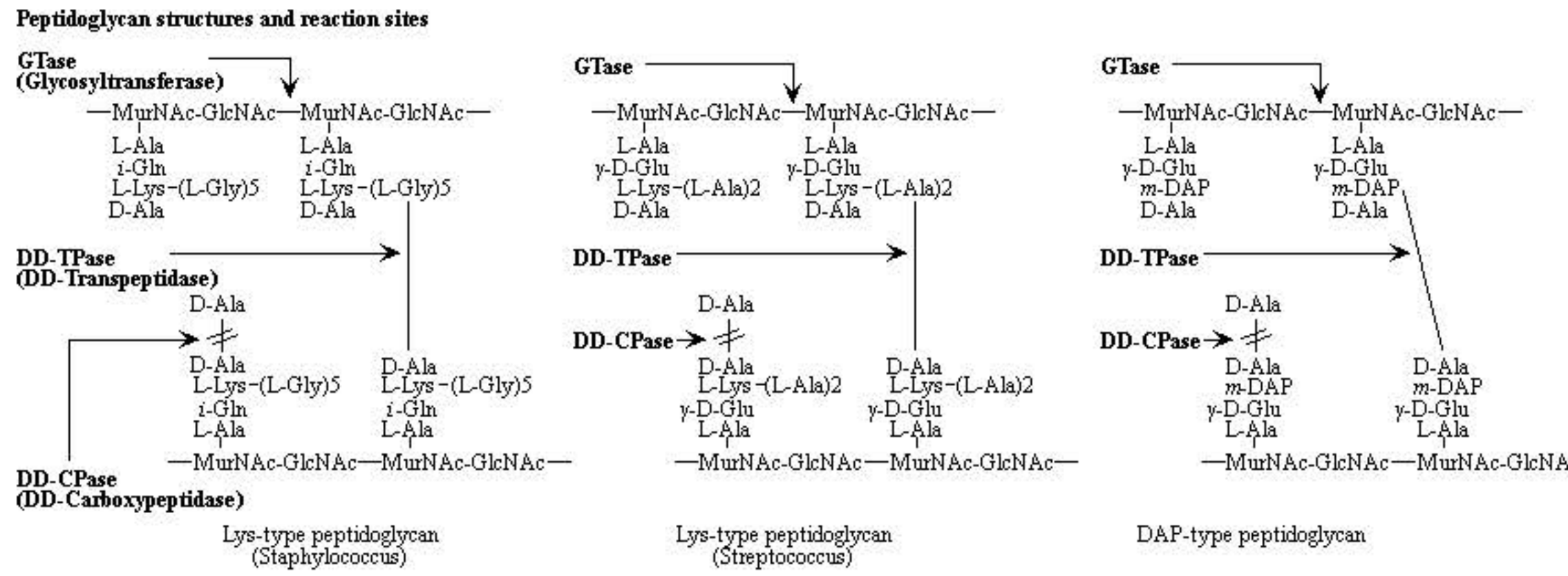
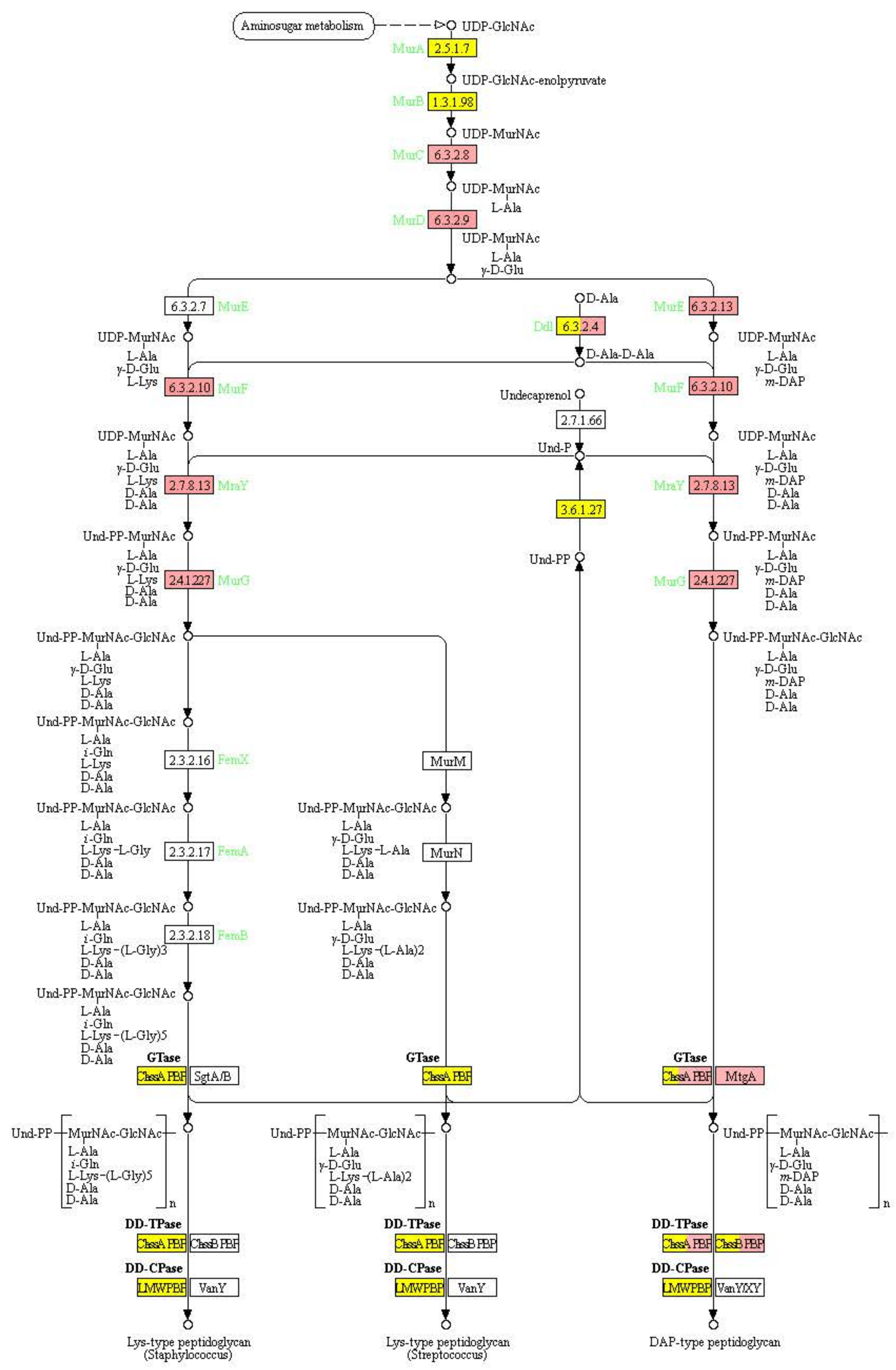
EF-Tu	S10	L3	L4	L23	L2	S19	L22	S3	RP-L16	L29	L7/L12 stalk					
	S20e	L3e	L4e	L23Ae	L8e	S15e	L17e	S3e		L35e						
S17	L14	L24		L5	S14	S8	L6			L18	S5	L30	L15	SecY		
S11e	L23e	L26e	S4e	L11e	S29e	S15Ae	L9e	L32e	L19e	L5e	S2e	L7e	L27Ae			
		IF1		L36	S13	S11	S4	RpoA		L17	L13	S9				
L34e	L14e			S18e		S14e	S9e	L18e		L13Ae	S16e					
EF-Tu,G	S7	S12		L7A	RpoC,B		L7/L12	L12	L10	L1	L11					
	S5e	S23e	L30e	L7Ae				LP1,LP2	LP0	L10Ae	L12e					
S2	EF-Ts	IF2		S15	IF3		L35	L20	L34	RF1		L31	L32	L9	S18	S6
SAe		S13e														
L28	L33	L21	L27	FtsY,Ffh		S16	L19	S1	S20	S21	L25					
L10e	L13e	L15e	L21e	L24e	L31e	L35Ae	L37e	L37Ae	L39e	L40e	L41e	L44e				
S3Ae	S6e	S8e	S17e	S19e	S24e	S25e	S26e	S27e	S27Ae	S28e	S30e	LX				
L6e	L18Ae	L22e	L27e	L28e	L29e	L36e	L38e									
S7e	S10e	S12e	S21e													

PURINE METABOLISM

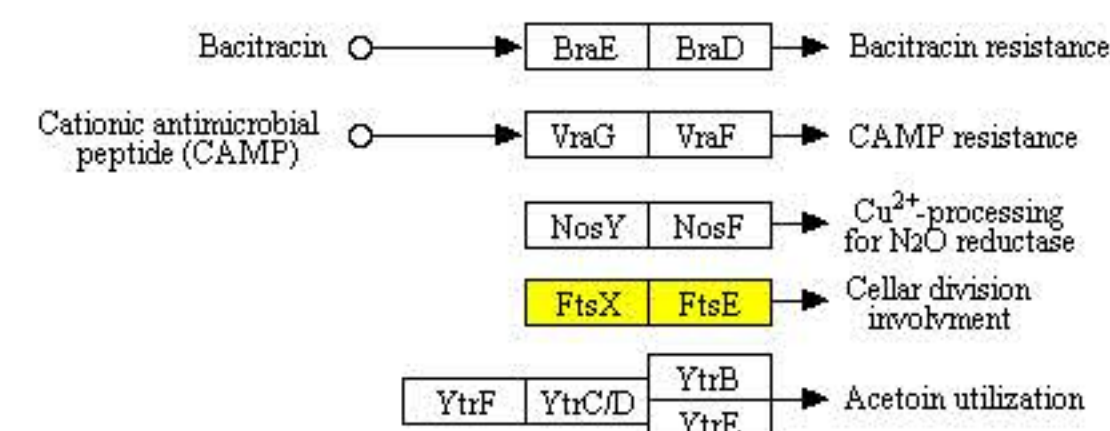
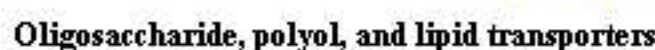


log₂(Fold Change)

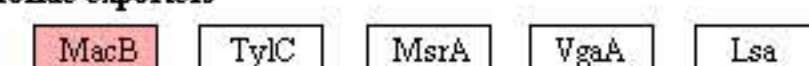
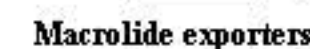
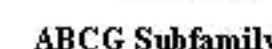
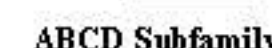
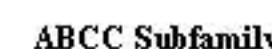
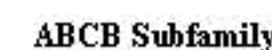




Mineral and organic ion transporters



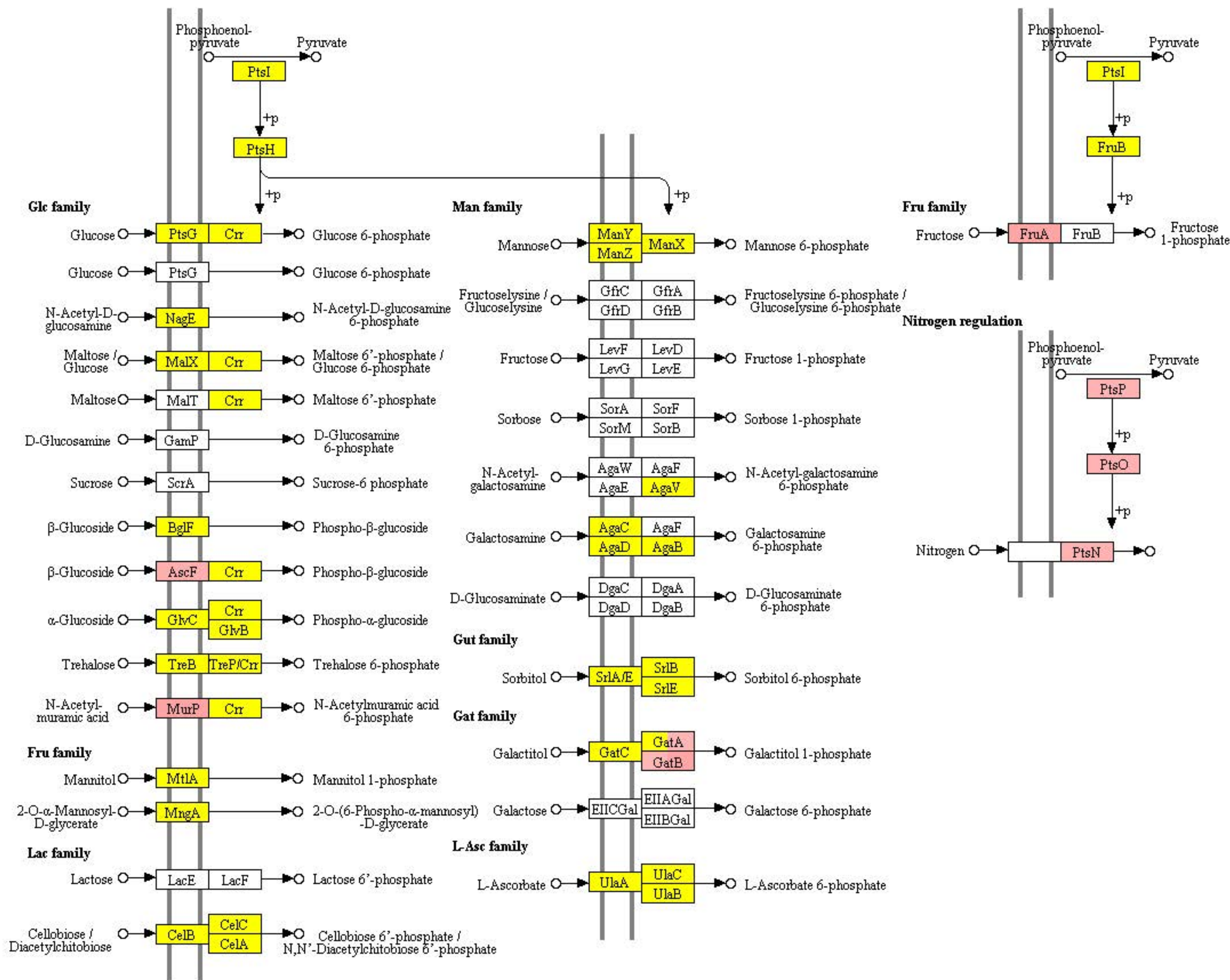
ABCA Subfamily



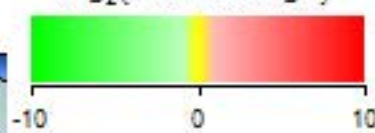
Other putative ABC transporters



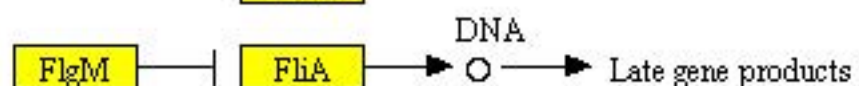
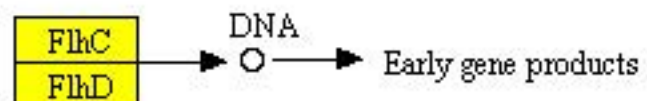
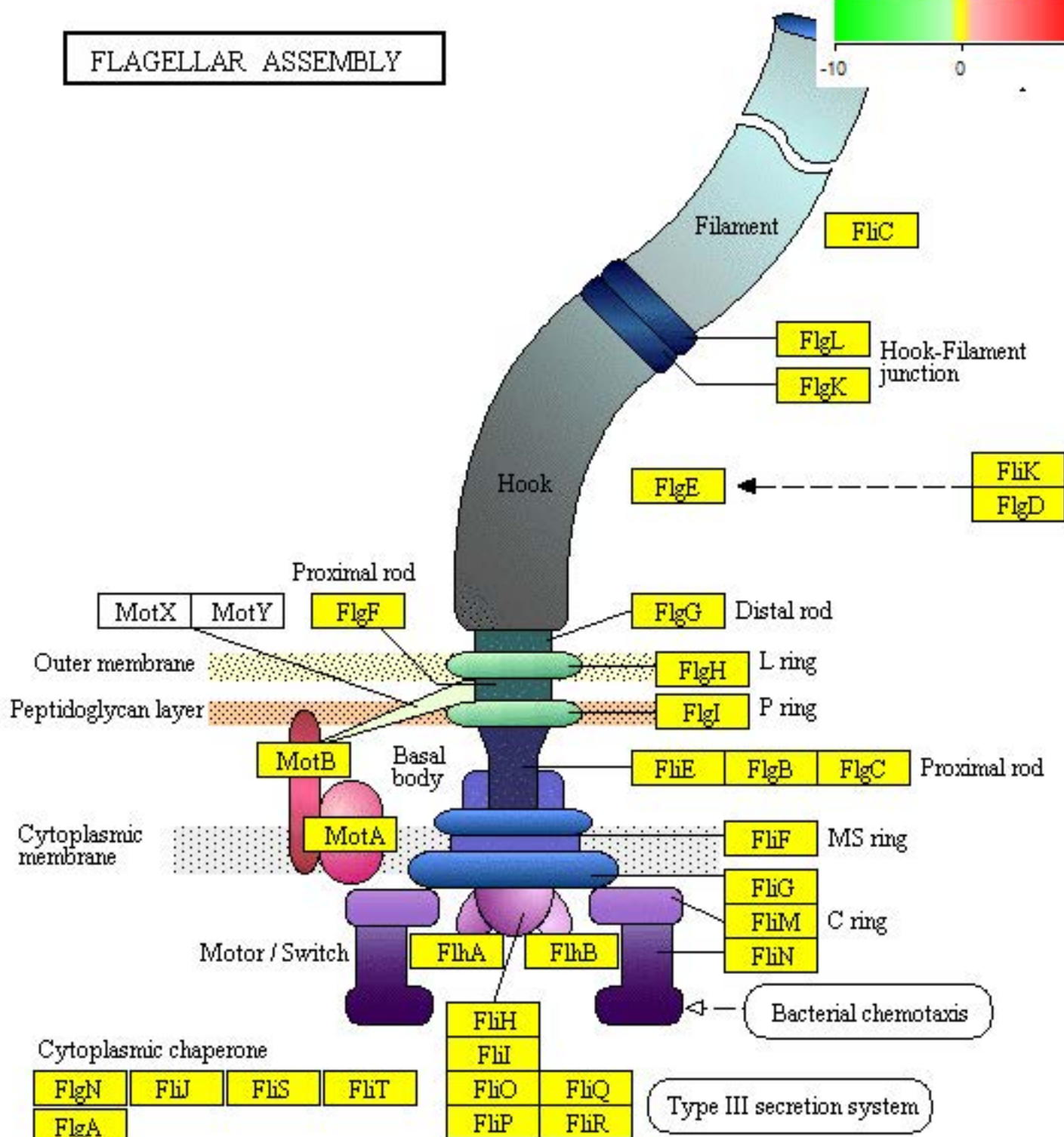
PHOSPHOTRANSFERASE SYSTEM (PTS)

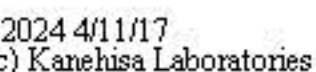


$\log_2(\text{Fold Change})$

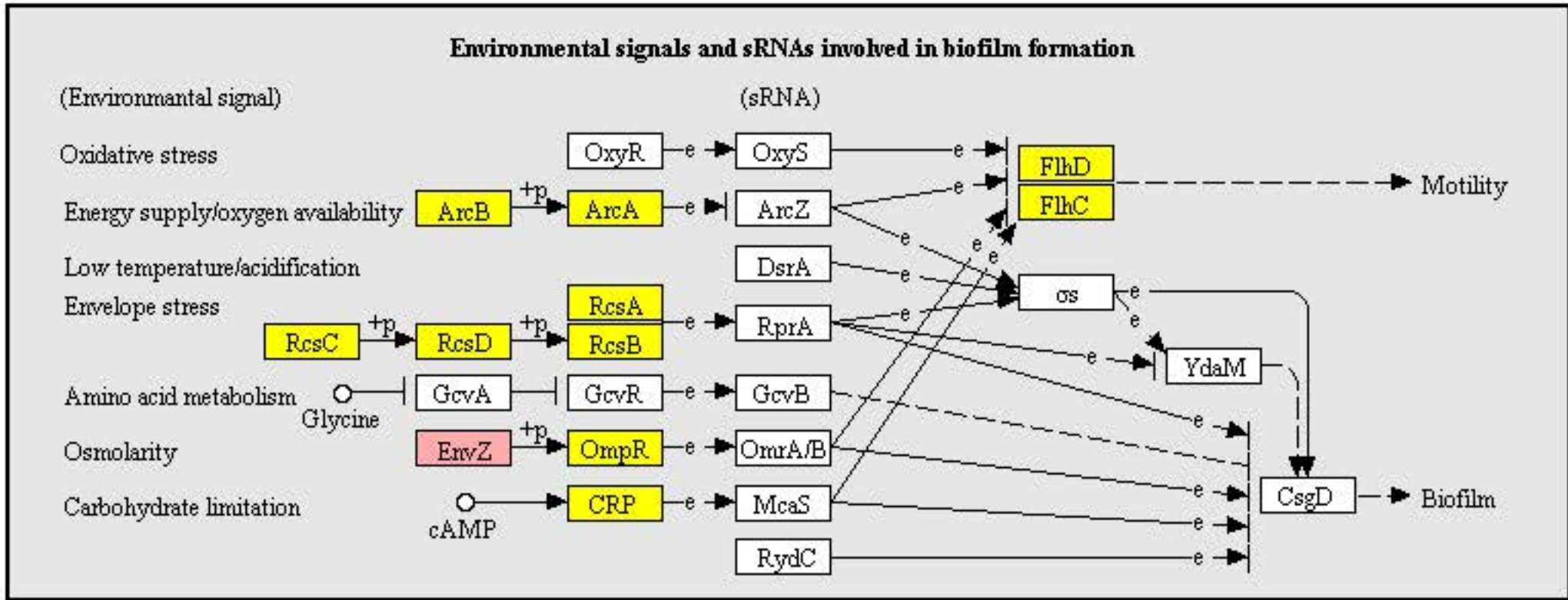
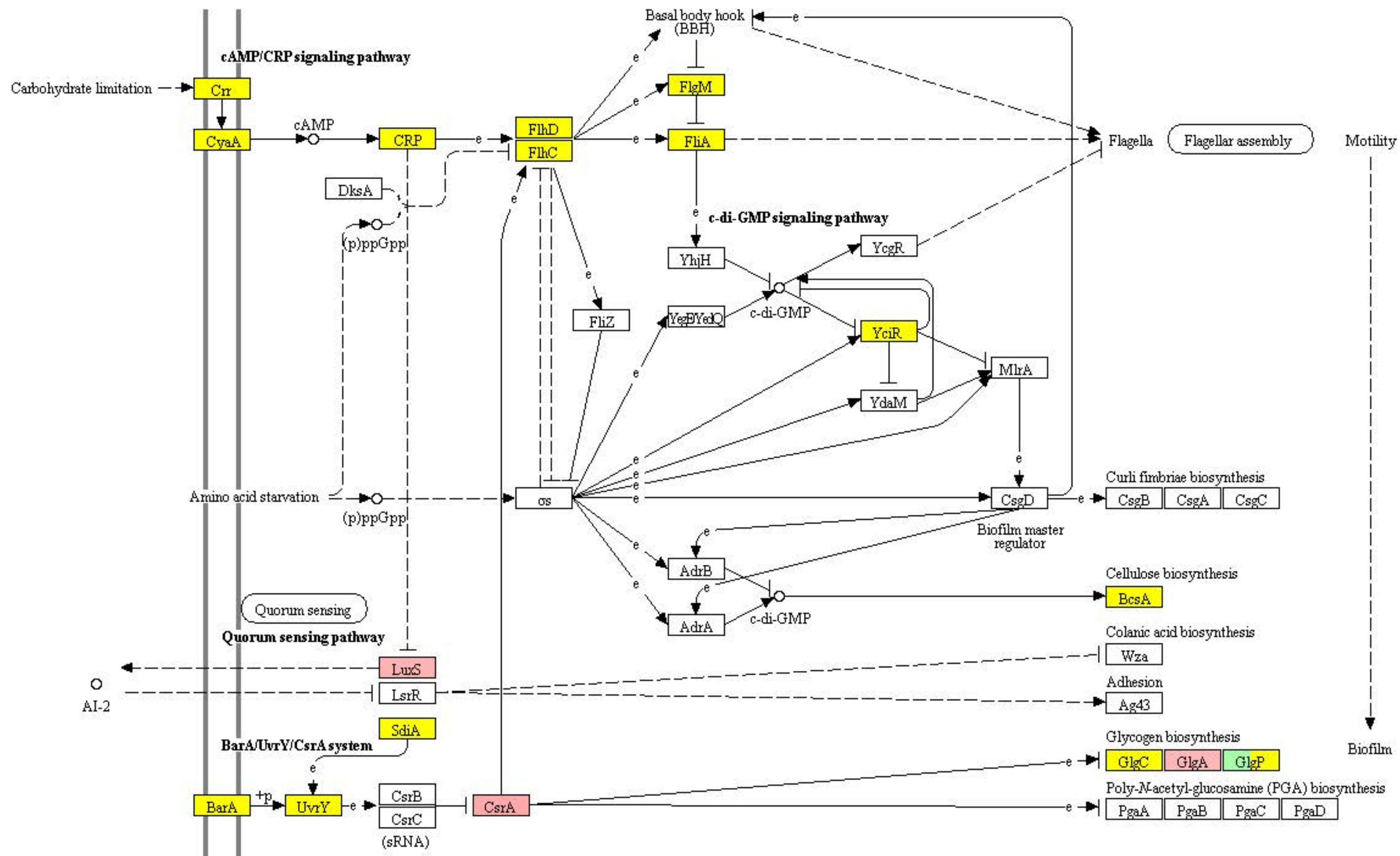
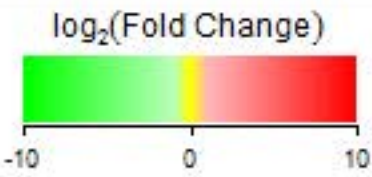


FLAGELLAR ASSEMBLY





BIOFILM FORMATION - ESCHERICHIA COLI



BACTERIAL CHEMOTAXIS

