MycoCosm KEGG Browser

KEGG stands for Kyoto Encyclopedia of Genes and Genomes at http://www.genome.jp/kegg/, which maintains a curated set of EC-annotated enzymes and their pathways. Each portal's KEGG Browser facilitates display and discovery of MycoCosm's KEGG-annotated genes.

Scenario: You have plated a variety of yeasts on a variety of carbon sources, and discovered that some members of the Pichiaceae grow on galactose (e.g. *Dekkera bruxellensis*) and some do not (e.g. *Pichia membranifaciens*). Use MycoCosm to find genes that could explain this metabolic difference.

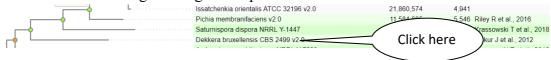
1) Go to the MycoCosm Pichiaceae PhyloGroup at genome.jgi.doe.gov/Pichiaceae :

П	REE	SEARCH	BLAST	ANNOTATIONS -	MCL CLUSTERS DO	OWNLOAD	INFO HELP!
G	rou	p Name:	I	Pichiaceae			
	##	Name			Assembly Length	# Genes	Published
	1	Candida arab	inoferment	ans NRRL YB-2248 v1	.0 13,234,078	5,861	<u>Riley R et al., 2016</u>
	2	Candida boid	inii NRRL Y	<u>Y-2332</u>	19,278,657	6,230	Krassowski T et al., 2018
	3	Dekkera brux	ellensis CE	<u>3S 2499 v2.0</u>	13,385,232	5,636	Piskur J et al., 2012
	4	Issatchenkia	orientalis A	TCC 32196 v2.0	21,860,574	4,941	
	5	Issatchenkia	orientalis A	TCC PTA-6658 v2.0	23,081,982	4,909	
	6	Issatchenkia	orientalis S	D108 v2.0	21,693,996	4,925	
	7	Nakazawaea	wickerham	iii NRRL Y-2563	10,857,285	5,490	Krassowski T et al., 2018
	8	Ogataea para	polymorph	a <u>DL-1</u>	8,874,589	5,325	
	9	Ogataea poly	morpha NC	CYC 495 leu1.1 v2.0	8,974,850	5,177	Riley R et al., 2016
	10	Pichia kudriav	zevii CBS	<u>573</u>	10,812,555	5,140	Douglass AP et al., 2018
	11	Pichia membr	ranifaciens	<u>v2.0</u>	11,584,685	5,546	Riley R et al., 2016
	12	Saturnispora	dispora NF	RRL Y-1447	10,120,266	4,881	Krassowski T et al., 2018

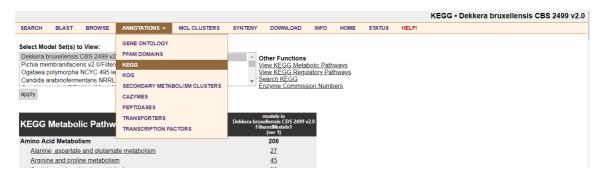
2) To verify that *Dekkera* (which grows on galactose) and *Pichia* (which does not) are sibling taxa, click on 'TREE':

				Tree • Pichiace
TREE SEARCH BLAST ANNOTATIONS -	MCL CLUSTERS DOWNLOA	D INFO HELP!		
To Default Expand All	Filter Clear To Root Re	load Support: ● 1.0 ● >=.70 ● <.70 ○ none		
	0.1	Name	Assembly Length (bp)	Genes Count Published
	r	Issatchenkia orientalis ATCC PTA-6658 v2.0	23,081,982	4,909
Pichia kudriavze	wii:4	Issatchenkia orientalis SD108 v2.0	21,693,996	4,925
Pichia:5		Pichia kudriavzevii CBS573	10,812,555	5,140 Douglass AP et al., 2018
	L	Issatchenkia orientalis ATCC 32196 v2.0	21,860,574	4,941
• L		Pichia membranifaciens v2.0	11,584,685	5,546 Riley R et al., 2016
L		Saturnispora dispora NRRL Y-1447	10,120,266	4,881 Krassowski T et al., 201
		Dekkera bruxellensis CBS 2499 v2.0	13,385,232	5,636 Piskur J et al., 2012
		Ambrosiozyma philentoma NRRL Y-7523	17,710,905	6,975 Krassowski T et al., 201
Pichiaceae:12 taea:3		Ogataea polymorpha NCYC 495 leu1.1 v2.0	8,974,850	5,177 Riley R et al., 2016
Picniaceae:12 laca.5		Ogataea parapolymorpha DL-1	8,874,589	5,325
T L		Candida arabinofermentans NRRL YB-2248 v1.0	13,234,078	5,861 Riley R et al., 2016
L		Candida boidinii NRRL Y-2332	19,278,657	6,230 Krassowski T et al., 2018

3) Click on 'Dekkera' to go to its genome portal:



4) Click on "ANNOTATIONS => KEGG" to go to the portal's KEGG browser:



5) Scroll down to the 'Carbohydrate Metabolism' section, and find the subsection 'Galactose metabolism'. Dekkera has 24 genes annotated to this metabolic pathway:

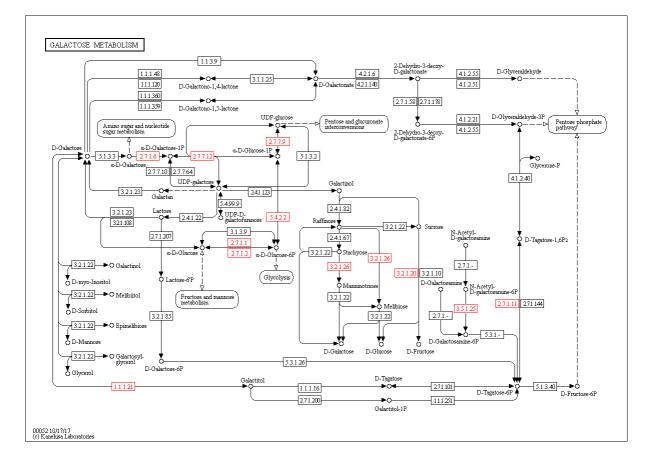
Carbohydrate Metabolism	332		
Amino sugar and nucleotide sugar metabolism	<u>68</u>		
Ascorbate and aldarate metabolism	<u>21</u>		
Butanoate metabolism	<u>34</u>		
C5-Branched dibasic acid metabolism	<u>2</u>		
Citrate cycle (TCA cycle)	<u>28</u>		
Fructose and mannose metabolism	<u>46</u>		
Galactose metabolism	<u>24</u>		
Glycolysis / Gluconeogenesis	<u>47</u>		
Glyoxylate and dicarboxylate metabolism	<u>10</u>		
Inositol phosphate metabolism	<u>27</u>		

- 6) Click on 'Galactose metabolism' to drill down into the KEGG hierarchy and list the EC numbers of that pathway.
- 7) Go to the 'Select Model Set(s) to View' list box and select *Dekkera* and *Pichia* and click the 'apply' button. The *Dekkera* and *Pichia* galactose metabolism gene counts are side-by-side and may be directly compared. Galactokinase (EC = 2.7.1.6) and UDPglucose-hexose-1-phosphate uridylyltransferase (2.7.7.12) are each present in *Dekkera* but not in *Pichia*:

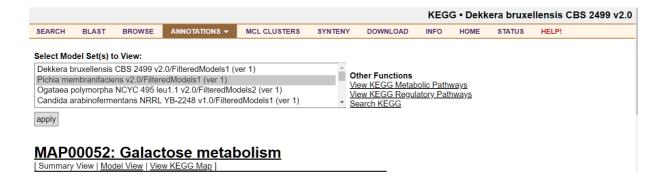
											KEGG • Dekkera bruxellensis CB	S 2499 v2.0
EARCH E	BLAST	BROWSE	ANNOTATIONS -	MCL CLUSTERS	SYNTENY	DOWNLOAD	INFO	HOME	STATUS	HELP!		
elect Model	Set(s) to	View:										
			2.0/FilteredModels1 (v redModels1 (ver 1)	ver 1)		ther Functions						
Ogataea poly	ymorpha	NCYC 495 le	au1.1 v2.0/FilteredMo		<u>\</u>	iew KEGG Meta iew KEGG Regi	bolic Path latory Pat	<u>ways</u> hwavs				
	binoferme	entans NRRL	YB-2248 v1.0/Filtere	dModels1 (ver 1)	* 3	earch KEGG						
apply												
		~ .										
			tose metak	olism								
C Number			<u></u>	Del	mode kkera bruxellens	is CBS 2499 v2.0 P	ichia memb	dels in ranifaciens v	/2.0 models in all selecte			
					FilteredN (ver		(v	dModels1 er 1)	model set			
	-	e 1-dehydro I 2-dehydrog	genase (NADP+)		0			0	0			
<u></u>		g			-				<u>.</u>		<u>.</u>	_
2.7.1.58	3	- 2-dehyd	ro-3-deoxygal	actonokinase)				0		0	0
<u>2.7.1.6</u>		galactok	inase						<u>1</u>		0	1
2.7.1.69	3	protein-l	Npi-phosphohi	stidinesug	ar phosp	hotransfera	ase		0		0	0
2.7.7.10)	UTPh	exose-1-phosi	ohate uridvlv	transfera	ISE			0		0	0
2.7.7.12	2	UDP-glu	icosehexose	e-1-phosphat	e uridyly	transferas	Э		1		0	1
2.7.7.9		UTPg	lucose-1-phos	phate uridyly	Itransfer	ase			2		2	4
3.1.1.25	5	1,4-lacto	onase						0		0	0
3.1.3.9		glucose-	-6-phosphatas	e					0		0	0
3.2.1.10		lactase							0		0	0
	_		ucosidase						6		2	8
3.2.1.20)	aibna-di										

- 8) Scroll back up to the now-familiar 'Select Model Set(s) to View' list box and select *Dekkera* only. Click 'apply' to show the *Dekkera* counts only.
- 9) Click 'View KEGG Map' to see a graphical display of the pathway. Only those enzyme boxes colored red are annotated as such in *Dekkera*. These include both 2.7.1.6(Galactokinase) and 2.7.7.12 (UDPglucose--hexose-1-phosphate uridylyltransferase):

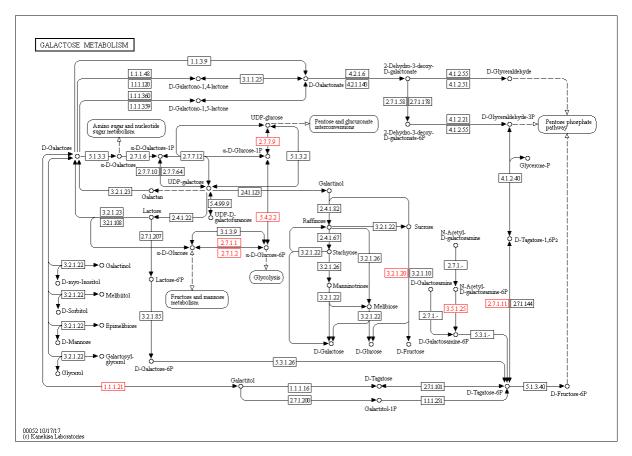




10) Use the web browser back button return to the now-familiar *Dekkera* galactose metabolism page and select *Pichia* only. Click 'apply' to show the *Pichia* counts only.



11) Click 'View KEGG Map' again, and again only those enzyme boxes colored red are annotated as such in *Pichia*. These include neither 2.7.1.6 nor 2.7.7.12. No wonder *Pichia* cannot grow on galactose!



References:

Riley R, Haridas S, Wolfe KH, Lopes MR, Hittinger CT, Göker M, Salamov AA, Wisecaver JH, Long TM, Calvey CH, Aerts AL, Barry KW, Choi C, Clum A, Coughlan AY, Deshpande S, Douglass AP, Hanson SJ, Klenk HP, LaButti KM, Lapidus A, Lindquist EA, Lipzen AM, Meier-Kolthoff JP, Ohm RA, Otillar RP, Pangilinan JL, Peng Y, Rokas A, Rosa CA, Scheuner C, Sibirny AA, Slot JC, Stielow JB, Sun H, Kurtzman CP, Blackwell M, Grigoriev IV, Jeffries TW. Comparative genomics of biotechnologically important yeasts. Proc Natl Acad Sci U S A. 2016 Aug 30;113(35):9882-7. doi: 10.1073/pnas.1603941113. Epub 2016 Aug 17. PubMed PMID: 27535936; PubMed Central PMCID: PMC5024638.