

Table 2: Representative metabolites decreased by antibiotic treatments

PATHWAY <sup>1</sup>	SUB-PATHWAY	BIOCHEMICAL NAME	Vanc 7d		Mix 7d		
			Ratio <sup>2</sup>	p-Value <sup>3</sup>	Ratio <sup>2</sup>	p-Value <sup>3</sup>	
Amino acid	Glycine, serine and threonine metabolism	glycine	0.76	0.013	0.85	0.126	
		dimethylglycine	0.61	< 0.001	0.66	< 0.001	
		serine	0.94	0.656	0.68	0.019	
		threonine	0.56	< 0.001	0.52	< 0.001	
		N-acetylthreonine	0.62	< 0.001	0.69	< 0.001	
	Alanine and aspartate metabolism	betaine	0.67	< 0.001	0.70	< 0.001	
		alanine	0.67	0.007	0.56	< 0.001	
		N-acetyl-beta-alanine	0.88	0.001	0.83	< 0.001	
		aspartate	0.63	0.005	1.00	0.912	
		N-acetylaspargate (NAA)	0.72	0.086	0.55	0.007	
	Glutamate metabolism	3-ureidopropionate	0.78	0.031	0.66	0.001	
		glutamate	0.65	0.002	0.61	< 0.001	
		glutamine	0.89	0.005	0.81	< 0.001	
	Phenylalanine & tyrosine metabolism	pyroglutamine*	0.36	< 0.001	0.31	< 0.001	
		phenyllactate (PLA)	1.69	< 0.001	0.14	< 0.001	
		phenylalanine	0.87	0.013	0.86	0.008	
		phenylacetate	0.78	0.046	0.84	0.137	
		p-cresol sulfate	0.09	< 0.001	0.11	< 0.001	
		tyrosine	0.95	0.193	0.88	0.003	
		3-(4-hydroxyphenyl)lactate	0.95	0.307	0.81	< 0.001	
		4-hydroxyphenylpyruvate	0.72	0.025	1.05	0.719	
		N-acetyltyrosine	0.93	0.367	0.82	0.013	
		phenylacetylglucine	0.19	< 0.001	0.93	0.942	
		2-(4-hydroxyphenyl)propionate	0.22	< 0.001	0.17	< 0.001	
		3-(3-hydroxyphenyl)propionate	0.54	0.045	0.35	0.024	
	Tryptophan metabolism	3-phenylpropionate (hydrocinnamate)	0.36	< 0.001	0.35	< 0.001	
		phenol sulfate	0.04	< 0.001	0.03	< 0.001	
		kynurenate	0.81	0.005	1.21	0.016	
		kynurenine	0.79	< 0.001	0.86	0.006	
		indoleacetate	0.54	< 0.001	0.64	0.003	
		serotonin (5HT)	3.11	< 0.001	0.85	0.657	
		N-acetyltryptophan	0.74	0.017	0.75	0.014	
		C-glycosyltryptophan*	0.87	0.004	0.89	0.009	
		3-indoxyl sulfate	0.66	0.077	0.47	0.001	
		indolepropionate	0.37	< 0.001	0.44	< 0.001	
		Valine, leucine and isoleucine metabolism	3-methyl-2-oxobutyrate	0.82	0.040	0.98	0.815
			3-methyl-2-oxovalerate	0.71	0.010	0.80	0.253
	beta-hydroxyisovalerate		1.10	0.327	1.51	0.007	
	isoleucine		0.81	0.003	0.91	0.101	
	leucine		0.84	< 0.001	0.92	0.033	
	N-acetylleucine		0.94	0.391	0.82	0.020	
	valine		0.92	0.040	1.01	0.807	
	2-hydroxyisobutyrate		0.72	< 0.001	0.63	< 0.001	
	3-hydroxyisobutyrate		0.70	0.007	0.81	0.179	
	alpha-hydroxyisovalerate		1.71	< 0.001	0.81	0.057	
isovalerylglucine	0.80		0.031	0.84	0.075		
2-methylbutyrylcarnitine	0.87		0.036	1.07	0.253		
Cysteine, methionine, SAM, taurine metabolism	methionine		0.84	< 0.001	0.81	< 0.001	
	N-acetylmethionine		0.81	< 0.001	0.72	< 0.001	
	2-hydroxybutyrate (AHB)		0.47	< 0.001	0.56	< 0.001	
Peptide	gamma-glutamyl	gamma-glutamylvaline	0.78	< 0.001	0.89	0.069	
		gamma-glutamylleucine	0.76	< 0.001	0.89	0.043	
		gamma-glutamylisoleucine*	0.80	0.002	0.80	0.002	
		gamma-glutamylmethionine	0.72	0.004	0.67	0.002	
		gamma-glutamylglutamine	0.74	< 0.001	0.70	< 0.001	
		gamma-glutamylphenylalanine	0.75	< 0.001	0.75	< 0.001	
		gamma-glutamyltyrosine	0.78	< 0.001	0.75	< 0.001	
		gamma-glutamylthreonine*	0.61	< 0.001	0.47	< 0.001	
		gamma-glutamylalanine	0.74	< 0.001	0.72	< 0.001	
		Carbohydrate	Fructose, mannose, galactose, starch, and sucrose metabolism	fructose	0.59	< 0.001	0.61
mannitol	0.92			0.843	0.06	< 0.001	
mannose	0.87			0.132	0.72	0.003	
Glycolysis, gluconeogenesis, pyruvate metabolism	1,5-anhydroglucitol (1,5-AG)		0.53	< 0.001	0.49	< 0.001	
	glycerate		0.69	0.046	0.81	0.195	
	glucose		0.71	0.003	0.67	< 0.001	
	lactate		0.63	0.005	1.39	0.037	
Energy	Krebs cycle	citrate	0.68	< 0.001	0.71	< 0.001	
		cis-aconitate	0.69	< 0.001	0.64	< 0.001	
		succinate	0.37	< 0.001	0.66	0.035	
		malate	0.48	< 0.001	0.58	< 0.001	
Nucleotide	Pyrimidine metabolism, cytidine containing	cytidine	0.88	0.054	0.81	0.003	
		5-methylcytidine	0.91	0.018	0.87	< 0.001	
		2'-deoxycytidine	0.94	0.010	0.89	< 0.001	
		uracil	0.67	0.012	0.99	0.999	
		uridine	0.81	< 0.001	0.83	< 0.001	
Cofactors and vitamins	Ascorbate and aldarate metabolism	pseudouridine	0.98	0.370	0.93	0.013	
		gulono-1,4-lactone	0.51	0.003	0.58	0.005	
		ascorbate (Vitamin C)	0.55	0.184	0.38	0.030	
Xenobiotics	Benzoate metabolism	threonate	0.55	< 0.001	0.42	< 0.001	
		hippurate	0.06	< 0.001	0.06	< 0.001	
		catechol sulfate	0.03	< 0.001	0.02	< 0.001	
	Food component/Plant	4-ethylphenylsulfate	0.08	< 0.001	0.08	< 0.001	
		indoleacrylate	0.43	< 0.001	0.48	< 0.001	
		cinnamate	0.48	< 0.001	0.35	< 0.001	
		stachydrine	0.59	< 0.001	0.62	< 0.001	
		homostachydrine*	0.47	< 0.001	0.50	< 0.001	
Lipid	Fatty acid, dicarboxylate	2-hydroxyglutarate	0.51	0.013	0.94	0.495	
		tetradecanedioate	0.81	0.032	0.97	0.861	
		hexadecanedioate	0.75	0.029	0.72	0.014	
	Bile acid metabolism	cholate	0.01	< 0.001	0.00	< 0.001	
		12-dehydrocholate	0.07	< 0.001	0.07	< 0.001	
		glycocholate	0.03	< 0.001	0.06	< 0.001	
		taurocholate	0.66	0.127	0.41	0.005	
		3-dehydrocholate	0.30	0.017	0.47	0.108	
		deoxycholate	0.14	< 0.001	0.21	0.003	
		taurodeoxycholate	0.42	< 0.001	0.50	0.001	
		beta-muricholate	0.01	< 0.001	0.04	< 0.001	
		alpha-muricholate	0.13	0.024	0.17	0.040	

<sup>1</sup> Comprehensive list of all pathways are shown in supplement table 1.

<sup>2</sup> Change reflected as ratio of 7 days (7d) relative to day 0 (0d, prior to antibiotic treatment). Vanc - vancomycin, Mix - mixed antibiotics.

<sup>3</sup> Significance indicated by repeated measure ANOVA test.