

“Plant-based diet index and metabolic risk in older men: exploring the role of the gut microbiome” Y Li et al. Online Supplemental Material

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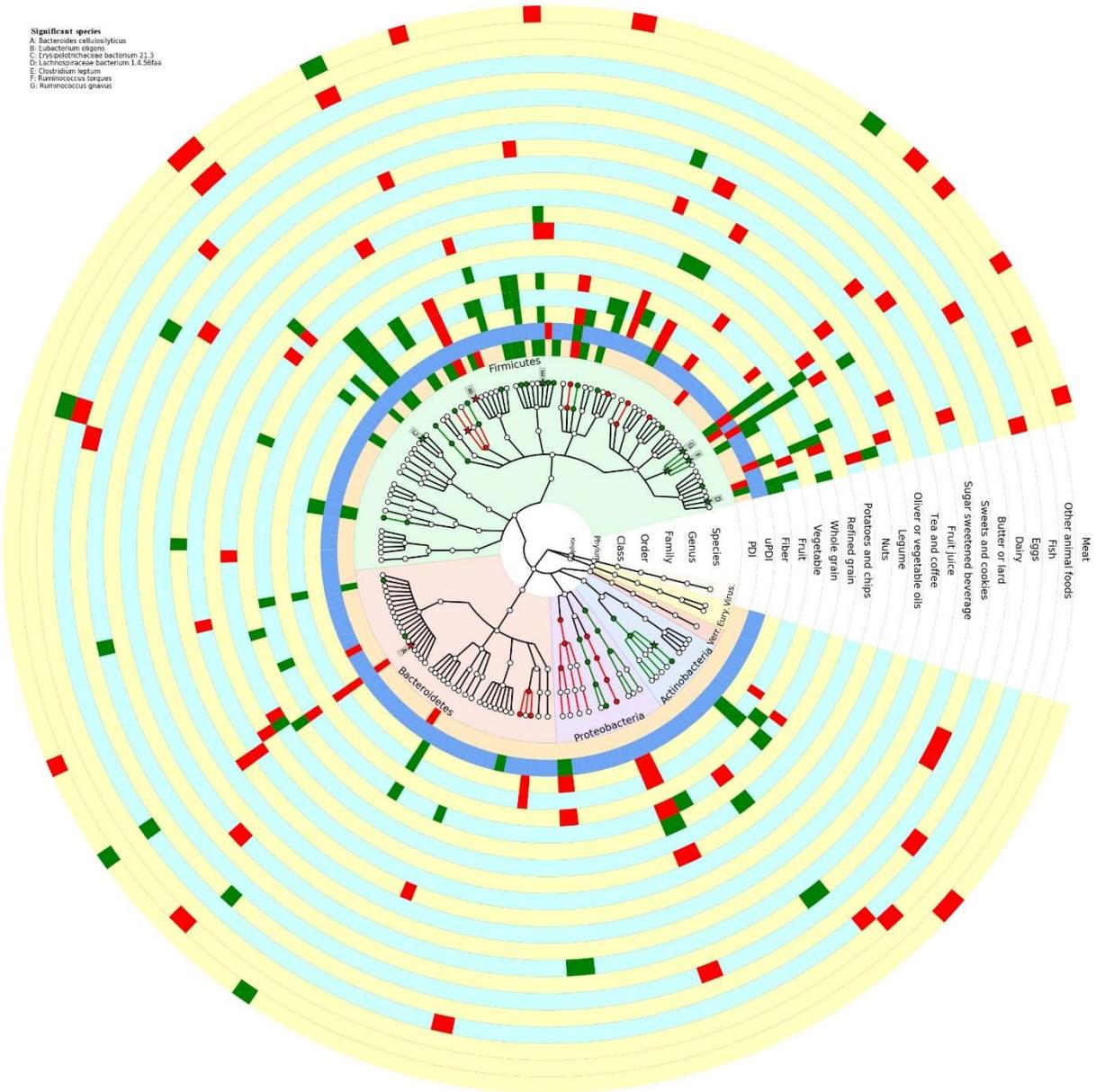
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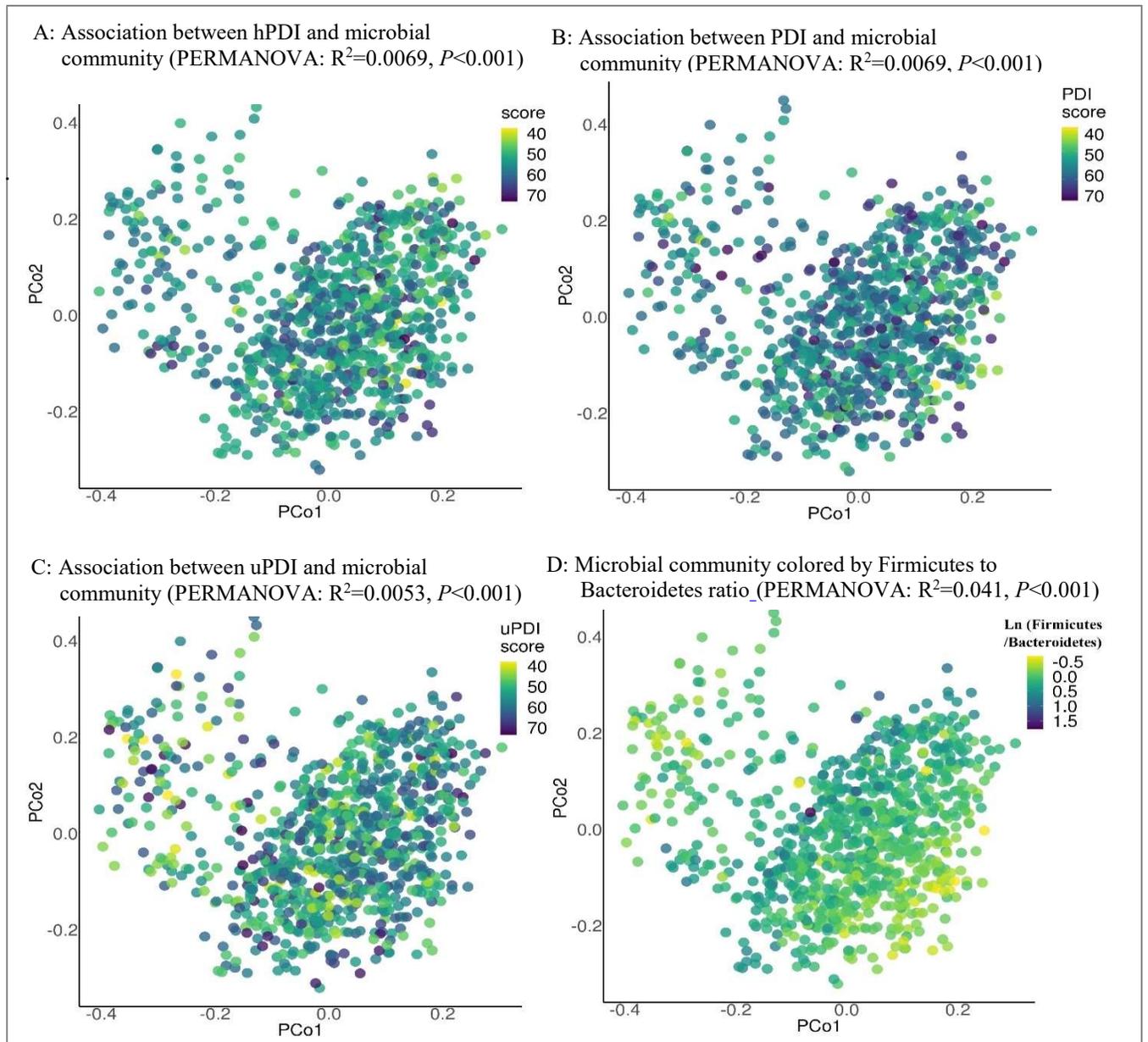
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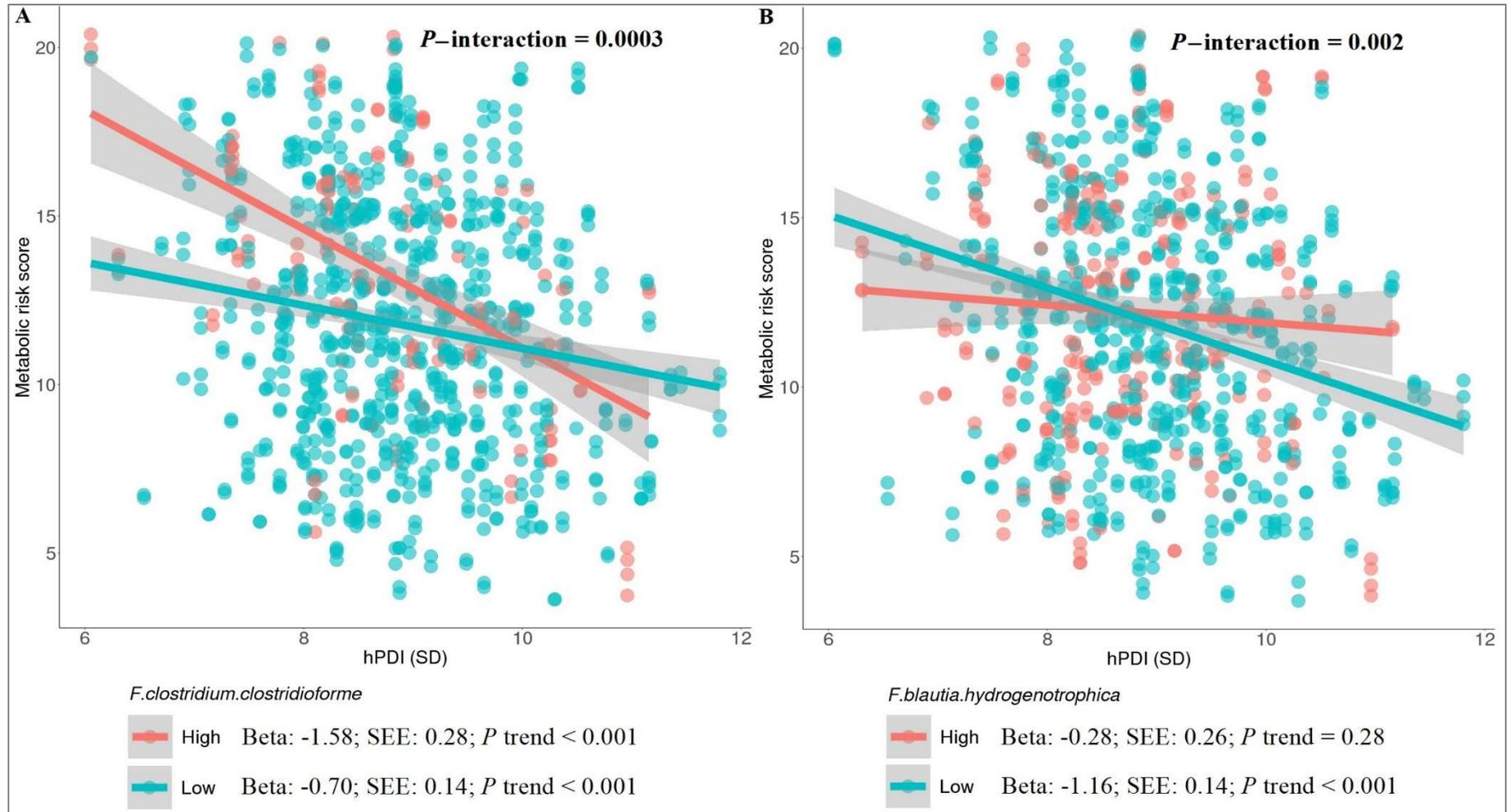


Supplemental Figure 3. Associations between PDIs and microbial communities based on 911 repeated measurements of the 303 participants.

Principal coordinates analysis based on Bray-Curtis dissimilarity were performed using taxonomic data at species level; the first 2 principal coordinates (PCo) were plotted by healthy plant-based diet index (hPDI) (A), PDI (B), unhealthy PDI (uPDI) (C), Firmicutes to Bacteroidetes ratio (D). A PERMANOVA test was used to evaluate the association of overall microbial compositions and PDIs, with multivariable models adjusted for repeated measurements (participants ID as random intercept), age, energy intake, alcohol, smoking, physical activity, using of antibiotics, consumed any probiotics and fecal sample characteristics (the Bristol stool chart).



Supplemental Figure 4. Interactions between healthy plant-based diet index (hPDI) and species on metabolic risk based on 911 repeated measurements of the 303 participants.



(SD: standard deviation; FDR for interaction: 0.01 (5A) and 0.036 (5B) after false discovery rate correction following the Benjamini–Hochberg).

Supplemental Table 1: List of enzymes that were significantly associated with hPDI

Enzymes	beta coefficients	SEM	N.not.0	P-value	Q-value	NAME of enzymes
ecd1_17_99_5	-0.001339601	0.000169	629	6.00E-15	3.31E-11	transferred entry: 1.17.98.1
ecd2_9_1_1	-0.001627706	0.000213	892	5.12E-14	1.42E-10	l-seryl-trna(sec) selenium transferase
ecd5_5_1_1	0.002007925	0.000276	852	8.06E-13	1.48E-09	muconate cycloisomerase
ecd3_6_3_2	-0.001072414	0.000154	705	5.97E-12	8.25E-09	magnesium-importing atpase
ecd1_1_1_2	-0.001642922	0.000253	809	1.33E-10	1.47E-07	alcohol dehydrogenase (nadp(+))
ecd5_4_99_62	-0.002074804	0.000325	838	2.91E-10	2.51E-07	d-ribose pyranase
ecd6_2_1_26	-0.002083701	0.000327	856	3.18E-10	2.51E-07	o-succinylbenzoate--coa ligase
ecd3_5_4_10	0.001829211	0.00029	895	4.55E-10	3.14E-07	Imp cyclohydrolase
ecd2_4_1_1	0.00294311	0.000486	895	2.04E-09	1.03E-06	glycogen phosphorylase
ecd3_2_1_49	-0.001009516	0.00017	661	4.34E-09	1.60E-06	alpha-n-acetylgalactosaminidase
ecd2_3_1_48	-0.000939905	0.00016	757	6.43E-09	2.22E-06	histone acetyltransferase
ecd3_6_5_3	0.001630807	0.000282	895	9.66E-09	3.14E-06	protein-synthesizing gtpase
ecd2_7_9_1	0.001076647	0.000189	894	1.80E-08	4.79E-06	pyruvate, phosphate dikinase
ecd4_4_1_8	-0.001453846	0.000267	810	6.60E-08	1.46E-05	cystathionine beta-lyase
ecd3_1_21_3	0.002178838	0.000405	894	9.69E-08	1.91E-05	type I site-specific deoxyribonuclease
ecd2_7_7_85	-0.001027477	0.000192	771	1.14E-07	2.17E-05	diadenylate cyclase
ecd3_1_4_16	-0.001044992	0.000207	589	5.67E-07	7.46E-05	2',3'-cyclic-nucleotide 2'-phosphodiesterase
ecd4_3_1_18	-0.001186432	0.000236	892	5.87E-07	7.55E-05	d-serine ammonia-lyase
ecd1_2_1_16	-0.00100538	0.000203	599	8.69E-07	0.000101552	succinate-semialdehyde dehydrogenase (nad(p)(+))
ecd2_1_2_3	0.001499589	0.000304	895	9.51E-07	0.000103212	phosphoribosylaminoimidazolecarboxamide formyltran
ecd1_1_1_157	0.001251811	0.000254	894	1.03E-06	0.000109746	3-hydroxybutyryl-coa dehydrogenase
ecd2_7_1_12	-0.001405965	0.000287	363	1.18E-06	0.00011494	gluconokinase
ecd3_1_1_1	0.000971876	0.000199	789	1.21E-06	0.00011494	carboxylesterase
ecd1_3_99_2	0.001294741	0.000267	895	1.44E-06	0.000130029	transferred entry: 1.3.8.1
ecd3_6_3_12	-0.00199246	0.000412	888	1.54E-06	0.000134952	potassium-transporting atpase
ecd4_3_1_15	-0.001015337	0.00021	869	1.62E-06	0.000140058	diaminopropionate ammonia-lyase
ecd2_4_1_281	0.001657555	0.000348	895	2.19E-06	0.000182986	4-o-beta-d-mannosyl-d-glucose phosphorylase

ecd2_1_1_79	-0.001462186	0.000307	780	2.26E-06	0.00018361	cyclopropane-fatty-acyl-phospholipid synthase
ecd4_1_99_18	-0.001506866	0.000317	895	2.39E-06	0.00018368	cyclic pyranopterin phosphate synthase
ecd3_1_3_3	-0.000847365	0.000179	894	2.50E-06	0.000186766	phosphoserine phosphatase
ecd1_11_1_6	-0.00165279	0.000357	895	4.28E-06	0.00027378	catalase
ecd2_6_1_42	0.001346271	0.000291	895	4.22E-06	0.00027378	branched-chain-amino-acid transaminase
ecd3_5_1_25	-0.001714207	0.00037	895	4.21E-06	0.00027378	n-acetylglucosamine-6-phosphate deacetylase
ecd2_1_1_176	-0.001049875	0.000228	315	4.80E-06	0.000298072	16s rrna (cytosine(967)-c(5))-methyltransferase
ecd3_6_3_30	0.001427097	0.000312	881	5.53E-06	0.000338382	fe(3+)-transporting atpase
ecd2_7_6_5	0.001374286	0.000303	895	6.47E-06	0.000376459	gtp diphosphokinase
ecd6_2_1_13	-0.001070014	0.000241	672	1.01E-05	0.000538135	acetate--coa ligase (adp-forming)
ecd2_3_1_89	-0.001166636	0.000265	877	1.19E-05	0.000618525	tetrahydrodipicolinate n-acetyltransferase
ecd4_1_3_6	0.001396489	0.000317	890	1.20E-05	0.000618525	citrate (pro-3s)-lyase
ecd1_2_1_12	-0.001304921	0.000301	882	1.66E-05	0.000779253	glyceraldehyde-3-phosphate dehydrogenase (phosphor
ecd2_7_1_107	-0.001728432	0.0004	888	1.71E-05	0.000779253	diacylglycerol kinase (atp)
ecd6_1_1_4	0.001911903	0.000441	895	1.60E-05	0.000779253	leucine--trna ligase
ecd6_3_4_6	0.000678436	0.000157	885	1.64E-05	0.000779253	urea carboxylase
ecd3_6_1_7	-0.001785306	0.000414	893	1.79E-05	0.000783982	cylphosphatase
ecd2_4_1_18	0.002161944	0.000506	895	2.12E-05	0.0009016	1,4-alpha-glucan branching enzyme
ecd4_3_1_7	-0.001240355	0.000291	887	2.19E-05	0.000908875	ethanolamine ammonia-lyase
ecd1_1_1_31	-0.000714563	0.000169	726	2.59E-05	0.001028541	3-hydroxyisobutyrate dehydrogenase
ecd3_4_19_3	-0.00108473	0.000257	876	2.61E-05	0.001028541	pyroglutamyl-peptidase I
ecd3_1_13_5	-0.000930905	0.00022	885	2.67E-05	0.001037379	ribonuclease d
ecd3_1_3_45	-0.001316062	0.000312	782	2.69E-05	0.001041083	3-deoxy-manno-octulosonate-8-phosphatase
ecd4_2_1_52	-0.000722193	0.000173	612	3.22E-05	0.001211667	transferred entry: 4.3.3.7
ecd1_1_99_33	-0.000960083	0.00023	282	3.31E-05	0.001235873	formate dehydrogenase (acceptor)
ecd2_4_2_3	-0.001316639	0.000318	894	3.78E-05	0.001370968	uridine phosphorylase
ecd1_16_3_1	-0.0016048	0.000391	881	4.52E-05	0.001562436	ferroxidase
ecd3_4_11_19	0.000994408	0.000244	850	4.94E-05	0.001667995	d-stereospecific aminopeptidase
ecd3_6_3_24	-0.001093757	0.000271	490	5.79E-05	0.001828481	nickel-transporting atpase
ecd3_5_3_6	-0.001037761	0.000258	891	6.17E-05	0.001915256	arginine deiminase

ecd3_1_1_31	-0.001150826	0.000286	881	6.22E-05	0.001920455	6-phosphogluconolactonase
ecd3_2_2_8	-0.000746684	0.000187	384	7.05E-05	0.002128083	ribosylpyrimidine nucleosidase
ecd1_1_1_272	-0.000812296	0.000204	869	7.26E-05	0.002157804	d-2-hydroxyacid dehydrogenase (nadp(+))
ecd3_1_1_61	0.001416032	0.000361	895	9.33E-05	0.002630463	protein-glutamate methylesterase
ecd2_1_1_151	-0.000796347	0.000203	827	9.49E-05	0.002642209	cobalt-factor II c(20)-methyltransferase
ecd2_1_1_166	-0.001239135	0.000317	831	0.000101433	0.002761169	23s rrna (uridine(2552)-2'-o)-methyltransferase
ecd1_1_5_3	-0.001791144	0.000459	844	0.000103566	0.002805427	glycerol-3-phosphate dehydrogenase
ecd2_7_1_5	-0.001076914	0.000276	895	0.000104697	0.002815351	rhamnukinase
ecd3_2_2_23	-0.00117051	0.000302	882	0.000112882	0.002998983	dna-formamidopyrimidine glycosylase
ecd2_7_7_6	0.002685797	0.000698	895	0.00012742	0.003274992	dna-directed rna polymerase
ecd5_4_99_16	0.001158048	0.000301	891	0.000131285	0.00331271	maltose alpha-d-glucosyltransferase
ecd3_6_3_34	-0.001386165	0.000362	895	0.000135725	0.003342299	Iron-chelate-transporting atpase
ecd3_5_1_44	0.001336545	0.000349	893	0.000138319	0.003382093	protein-glutamine glutaminase
ecd1_1_1_159	-0.001766627	0.000463	893	0.000144669	0.003496755	7-alpha-hydroxysteroid dehydrogenase
ecd6_1_1_13	-0.001526742	0.000401	889	0.000148704	0.003546755	d-alanine--poly(phosphoribitol) ligase
ecd1_1_1_6	-0.001164777	0.000306	887	0.000149829	0.003553466	glycerol dehydrogenase
ecd1_17_4_2	0.000787679	0.000207	894	0.000152103	0.003591965	ribonucleoside-triphosphate reductase
ecd1_21_4_2	-0.000597738	0.000157	700	0.000153966	0.003620492	glycine reductase
ecd2_7_1_45	0.001276593	0.000337	895	0.000159651	0.003691355	2-dehydro-3-deoxygluconokinase
ecd3_6_3_44	-0.001435626	0.000381	836	0.000176106	0.003988366	xenobiotic-transporting atpase
ecd3_5_3_9	0.000735776	0.000195	835	0.000177496	0.004003439	allantoate deiminase
ecd3_1_3_25	-0.001110089	0.000298	893	0.000205089	0.004497316	Inositol-phosphate phosphatase
ecd5_1_3_14	-0.00103064	0.000277	895	0.000215434	0.004705486	udp-n-acetylglucosamine 2-epimerase (non-hydrolyzi
ecd1_21_4_4	-0.00055115	0.000149	653	0.000219078	0.004733685	betaine reductase
ecd1_4_1_21	0.000840911	0.000227	892	0.000218561	0.004733685	aspartate dehydrogenase
ecd3_1_6_1	-0.001158808	0.000315	633	0.000248262	0.005157505	arylsulfatase
ecd2_7_7_87	0.001194134	0.000325	895	0.000253147	0.00521975	l-threonylcarbamoyladenylate synthase
ecd5_3_1_25	-0.001230989	0.000336	895	0.0002647	0.00529975	l-fucose Isomerase
ecd2_1_1_107	-0.001002465	0.000277	893	0.000307926	0.005867587	uroporphyrinogen-iii c-methyltransferase
ecd2_4_1_7	-0.000964774	0.000267	665	0.000313028	0.00594431	sucrose phosphorylase

ecd1_15_1_2	0.001048577	0.00029	874	0.000318269	0.006019838	superoxide reductase
ecd1_1_1_251	-0.000747083	0.000209	868	0.000367481	0.006724166	galactitol-1-phosphate 5-dehydrogenase
ecd1_3_1_74	0.001254806	0.000354	895	0.000416699	0.007306555	2-alkenal reductase (nad(p)(+))
ecd3_4_15_5	-0.000623856	0.000176	521	0.00041782	0.007306555	peptidyl-dipeptidase dcp
ecd3_6_1_3	0.001238454	0.000351	894	0.000437682	0.007629742	adenosinetriphosphatase
ecd4_3_1_1	-0.001417611	0.000405	880	0.000487744	0.008293147	aspartate ammonia-lyase
ecd2_1_1_74	0.001407353	0.000404	894	0.000511295	0.008614082	(fadh(2)-oxidizing)
ecd2_7_7_77	-0.000927916	0.000266	890	0.00051889	0.008689046	molybdenum cofactor guanylyltransferase
ecd2_8_3_12	-0.000605121	0.000175	842	0.000587527	0.009577213	glutaconate coa-transferase
ecd5_3_1_13	-0.001165274	0.000339	895	0.000616179	0.00984105	arabinose-5-phosphate Isomerase
ecd3_5_2_17	-0.001366408	0.000399	374	0.000636749	0.00996792	hydroxyisourate hydrolase
ecd1_1_1_38	0.000904787	0.000265	894	0.000653691	0.010204222	malate dehydrogenase (oxaloacetate-decarboxylating)
ecd1_3_99_1	0.002018226	0.00059	895	0.000655826	0.01020872	deleted entry
ecd3_1_2_6	-0.001165853	0.000342	563	0.00068703	0.010516698	hydroxyacylglutathione hydrolase
ecd1_3_8_1	-0.001155643	0.000341	711	0.000726418	0.010917572	short-chain acyl-coa dehydrogenase
ecd3_2_1_86	-0.001460251	0.000432	895	0.000757458	0.011044093	6-phospho-beta-glucosidase
ecd3_2_1_4	0.001924385	0.00057	895	0.000767712	0.011150023	cellulase
ecd1_1_1_26	-0.000638398	0.000191	492	0.000855731	0.012001954	glyoxylate reductase
ecd2_8_3_8	-0.000871462	0.000261	895	0.000862767	0.012070006	acetate coa-transferase
ecd3_6_3_21	0.000979711	0.000294	894	0.000912893	0.012674985	polar-amino-acid-transporting atpase
ecd3_8_1_2	0.000854311	0.00026	893	0.001042334	0.014082978	(s)-2-haloacid dehalogenase
ecd3_1_1_11	0.002007398	0.000613	895	0.001096514	0.014565717	pectinesterase
ecd3_6_3_8	-0.000798854	0.000246	886	0.001222071	0.015680244	calcium-transporting atpase
ecd6_3_2_5	0.000857183	0.000264	895	0.001231779	0.015720117	phosphopantothenate--cysteine ligase
ecd3_2_1_8	0.00178968	0.000556	895	0.001330327	0.016557177	endo-1,4-beta-xylanase
ecd2_7_13_3	-0.002015436	0.000627	895	0.001365132	0.016672229	histidine kinase
ecd2_4_1_211	0.000896546	0.000282	875	0.001517642	0.018163874	1,3-beta-galactosyl-n-acetylhexosamine phosphoryla
ecd3_4_17_19	0.000792474	0.00025	890	0.00158763	0.018666471	carboxypeptidase taq
ecd5_1_3_11	0.00157484	0.000502	895	0.001753665	0.020220525	cellobiose epimerase
ecd2_3_1_128	-0.001216767	0.000391	895	0.001919876	0.021389582	ribosomal-protein-alanine n-acetyltransferase

ecd4_1_2_17	-0.000944583	0.000305	882	0.002034481	0.02200106	l-fucose-phosphate aldolase
ecd1_3_1_12	0.000850127	0.000276	894	0.002101027	0.022544227	prephenate dehydrogenase
ecd5_1_3_6	-0.000520925	0.00017	467	0.002216643	0.023556098	udp-glucuronate 4-epimerase
ecd6_3_4_14	-0.000805112	0.000263	895	0.002287875	0.02417361	biotin carboxylase
ecd3_2_1_89	0.00158396	0.000519	895	0.002336418	0.024535055	arabinogalactan endo-beta-1,4-galactanase
ecd3_4_13_19	0.000797005	0.000262	882	0.002415001	0.024944478	membrane dipeptidase
ecd2_7_1_31	-0.001026607	0.000338	895	0.00242778	0.025029686	glycerate 3-kinase
ecd2_5_1_30	0.000999697	0.000333	735	0.002748099	0.027461455	heptaprenyl diphosphate synthase
ecd2_4_99_16	0.000897152	0.0003	521	0.002875358	0.028424378	starch synthase (maltosyl-transferring)
ecd3_5_4_3	-0.000573124	0.000193	841	0.002991511	0.029104028	guanine deaminase
ecd1_1_1_58	0.000827581	0.000278	895	0.003011873	0.029235752	tagaturonate reductase
ecd3_5_3_1	-0.000643433	0.000218	881	0.003242734	0.030895424	arginase
ecd1_1_1_103	-0.001024031	0.000348	895	0.003303649	0.031367637	l-threonine 3-dehydrogenase
ecd1_5_3_1	-0.001071485	0.000364	887	0.00334612	0.031500275	sarcosine oxidase
ecd4_1_2_13	0.000914837	0.000313	895	0.003526845	0.032589746	fructose-bisphosphate aldolase
ecd3_5_1_47	-0.000755935	0.000259	788	0.003665186	0.033467475	n-acetyldiaminopimelate deacetylase
ecd2_7_1_51	-0.001010822	0.000348	853	0.00372115	0.033724276	l-fuculokinase
ecd2_3_1_8	0.000705309	0.000243	895	0.003808391	0.034376741	phosphate acetyltransferase
ecd3_1_3_11	-0.001084264	0.000375	895	0.003898333	0.034797809	fructose-bisphosphatase
ecd3_2_1_96	-0.00054227	0.000189	418	0.004143279	0.036445559	mannosyl-glycoprotein endo-beta-n-acetylglucosamin
ecd6_3_2_2	-0.001466335	0.000511	859	0.004243456	0.03704477	glutamate--cysteine ligase
ecd2_1_1_63	-0.001148331	0.000401	895	0.004329591	0.037626999	methylated-dna--[protein]-cysteine s-methyltransfe
ecd2_7_1_60	-0.000632056	0.000222	812	0.004432509	0.038071475	n-acylmannosamine kinase
ecd5_3_1_5	0.001080138	0.000379	895	0.004427119	0.038071475	xylose Isomerase
ecd1_14_13_81	0.000428179	0.000151	890	0.004565698	0.038935257	magnesium-protoporphyrin IX monomethyl ester (oxid
ecd3_2_1_85	-0.000825607	0.000291	807	0.00460722	0.039168462	6-phospho-beta-galactosidase
ecd4_2_2_n2	-0.000754545	0.000267	246	0.004759672	0.040278632	peptidoglycan lytic endotransglycosylase
ecd2_8_3_11	0.000506325	0.00018	369	0.004951548	0.041084466	citramalate coa-transferase
ecd5_3_1_4	0.001018079	0.000362	895	0.00507678	0.041749888	l-arabinose Isomerase
ecd1_2_99_7	-0.000799985	0.000285	871	0.005140411	0.042074045	aldehyde dehydrogenase (fad-independent)

ecd1_6_99_5	-0.001120732	0.0004	866	0.005171311	0.042210728	transferred entry: 1.6.5.11
ecd1_2_1_2	-0.000955516	0.000342	795	0.005305527	0.042738104	formate dehydrogenase
ecd1_3_5_2	-0.001017997	0.000367	893	0.00559278	0.044214163	dihydroorotate dehydrogenase (quinone)
ecd1_13_12_16	0.000629111	0.000227	738	0.005799846	0.045396525	nitronate monooxygenase
ecd3_6_3_41	-0.000605352	0.00022	782	0.00596647	0.046242232	heme-transporting atpase
ecd2_3_1_12	-0.000555552	0.000202	501	0.00616655	0.047022603	dihydrolipoyllysine-residue acetyltransferase
ecd3_6_1_1	0.001016092	0.000371	895	0.006234823	0.047196757	Inorganic diphosphatase
ecd5_3_1_26	-0.000718956	0.000264	539	0.006599837	0.049218214	galactose-6-phosphate Isomerase
ecd1_1_1_57	0.000781443	0.000289	894	0.007006703	0.051106777	fructuronate reductase
ecd4_1_3_27	-0.001027838	0.000381	894	0.007150324	0.051785965	anthranilate synthase
ecd2_5_1_17	-0.000776686	0.000289	893	0.007247655	0.052285305	cob(i)yrinic acid a,c-diamide adenosyltransferase
ecd4_1_1_3	0.001021565	0.00038	895	0.007280813	0.052456027	oxaloacetate decarboxylase
ecd3_6_4_12	0.001503723	0.000563	895	0.007666712	0.054315708	dna helicase
ecd3_1_1_47	-0.000581331	0.000218	742	0.007827627	0.054892724	1-alkyl-2-acetyl-glycerophosphocholine esterase
ecd2_1_1_242	-0.000720949	0.000272	870	0.008095847	0.056132559	16s rrna (guanine(1516)-n(2))-methyltransferase
ecd1_3_1_44	0.000856328	0.000323	873	0.008128052	0.056227332	trans-2-enoyl-coa reductase (nad(+))
ecd2_1_2_5	-0.001122688	0.000423	794	0.008129866	0.056227332	glutamate formimidoyltransferase
ecd3_2_1_21	0.001131331	0.000427	895	0.008222824	0.056672081	beta-glucosidase
ecd4_1_1_17	-0.000703879	0.000266	536	0.008267347	0.056822588	ornithine decarboxylase
ecd1_1_1_40	-0.000735	0.000278	793	0.008312493	0.057049174	malate dehydrogenase (oxaloacetate-decarboxylating)
ecd3_5_4_5	-0.00065621	0.000249	895	0.008446963	0.057414409	cytidine deaminase
ecd3_4_23_43	0.000947807	0.000361	876	0.008772226	0.058910665	prepilin peptidase
ecd1_8_1_2	-0.001055718	0.000406	588	0.009542513	0.06270146	assimilatory sulfite reductase (nadph)
ecd2_3_1_117	-0.000816453	0.000318	881	0.010355782	0.066598981	2,3,4,5-tetrahydropyridine-2,6-dicarboxylate n-suc
ecd3_2_1_94	-0.000574652	0.000224	298	0.010508324	0.067131788	glucan 1,6-alpha-isomaltosidase
ecd2_7_1_4	0.000945242	0.000369	895	0.010600035	0.067405974	fructokinase
ecd3_2_1_18	-0.000853757	0.000335	811	0.01096525	0.069250254	exo-alpha-sialidase
ecd1_8_1_4	-0.001266765	0.000498	895	0.011094033	0.069903792	dihydrolipoyl dehydrogenase
ecd4_2_2_8	-0.000794777	0.000314	234	0.011572977	0.071725735	heparin-sulfate lyase
ecd5_3_3_3	-0.000420878	0.000166	613	0.011643626	0.071971672	vinylacetyl-coa delta-isomerase

ecd4_2_1_32	0.000932529	0.00037	895	0.011821459	0.072826512	l(+)-tartrate dehydratase
ecd3_5_4_12	0.000761737	0.000303	894	0.012063129	0.073821539	dcmp deaminase
ecd3_1_4_53	-0.000622453	0.000248	570	0.012097218	0.073880369	3',5'-cyclic-amp phosphodiesterase
ecd6_6_1_2	-0.000470946	0.000188	607	0.012258303	0.074520775	cobaltochelataase
ecd5_3_3_10	-0.000640285	0.000256	174	0.012484643	0.075316744	5-carboxymethyl-2-hydroxymuconate delta-isomerase
ecd5_1_99_1	-0.00093288	0.000374	870	0.01278776	0.076477451	methylmalonyl-coa epimerase
ecd4_1_1_70	0.000709224	0.000285	895	0.012903607	0.076906123	glutaconyl-coa decarboxylase
ecd1_1_1_21	-0.000445408	0.000179	786	0.013079991	0.07755368	ldehyde reductase
ecd6_2_1_22	0.000637262	0.000257	868	0.013227157	0.078091099	[citrate (pro-3s)-lyase] ligase
ecd1_8_98_1	-0.000651324	0.000262	882	0.013242161	0.078096247	cob--com heterodisulfide reductase
ecd3_6_4_13	-0.001404777	0.000567	811	0.013343513	0.07833068	rna helicase
ecd3_2_1_25	0.000522769	0.000212	831	0.013966777	0.080506911	beta-mannosidase
ecd3_2_2_1	-0.000515195	0.000209	831	0.014024818	0.080562521	purine nucleosidase
ecd4_1_99_12	-0.000866351	0.000353	895	0.014175614	0.081089135	3,4-dihydroxy-2-butanone-4-phosphate synthase
ecd1_1_1_100	-0.001076763	0.000443	895	0.015265502	0.085295412	3-oxoacyl-[acyl-carrier-protein] reductase
ecd3_5_4_25	-0.000610163	0.000251	852	0.015333165	0.085586938	gtp cyclohydrolase II
ecd6_3_5_11	-0.000610376	0.000253	894	0.015836201	0.087862298	cobyrinate a,c-diamide synthase (glutamine-hydroly
ecd1_2_7_1	0.001034533	0.00043	890	0.016296427	0.089606024	pyruvate synthase
ecd1_2_4_1	-0.000483295	0.000201	481	0.016520266	0.090420998	pyruvate dehydrogenase (acetyl-transferring)
ecd5_1_1_11	0.000458182	0.000191	583	0.016526458	0.090420998	phenylalanine racemase (atp-hydrolyzing)
ecd4_1_1_31	-0.001129794	0.000472	557	0.016823638	0.091503372	phosphoenolpyruvate carboxylase
ecd2_5_1_1	-0.000483488	0.000203	381	0.017208067	0.092961207	dimethylallyltranstransferase
ecd2_3_1_222	-0.000804125	0.000338	895	0.017452867	0.093726477	phosphate propanoyltransferase
ecd3_2_1_n1	-0.000987846	0.000415	892	0.017584969	0.09434421	blood group b branched chain alpha-1,3-galactosida
ecd1_17_1_4	-0.000597617	0.000253	890	0.018408609	0.096789697	xanthine dehydrogenase
ecd1_6_5_5	-0.000616136	0.000261	624	0.018636273	0.097652587	nadph:quinone reductase
ecd3_1_3_27	-0.001052837	0.000447	880	0.018747824	0.098106511	phosphatidylglycerophosphatase
ecd3_4_25_2	-0.000885943	0.000378	750	0.019218849	0.100108567	hslu--hslv peptidase
ecd3_4_19_5	-0.000638517	0.000274	496	0.019795991	0.102331753	beta-aspartyl-peptidase
ecd2_1_1_113	-0.000664962	0.000287	762	0.020711638	0.105974546	site-specific dna-methyltransferase (cytosine-n(4)

ecd3_4_14_11	-0.00045004	0.000194	854	0.020862095	0.106214416	xaa-pro dipeptidyl-peptidase
ecd2_2_1_1	0.000895921	0.000387	895	0.02099664	0.106642859	transketolase
ecd1_1_1_69	0.000953466	0.000413	895	0.021037688	0.106704199	gluconate 5-dehydrogenase
ecd3_2_1_131	0.000606475	0.000263	871	0.021479347	0.107513472	xylan alpha-1,2-glucuronosidase
ecd3_5_2_10	-0.000571831	0.00025	602	0.022238646	0.110314863	creatininase
ecd1_2_7_8	0.001019056	0.000445	895	0.022314504	0.110591883	Indolepyruvate ferredoxin oxidoreductase
ecd1_1_1_67	0.000552144	0.000242	828	0.02274159	0.112000844	mannitol 2-dehydrogenase
ecd3_4_17_11	0.00033585	0.000147	583	0.022760939	0.112000844	glutamate carboxypeptidase
ecd3_2_1_151	0.000694082	0.000305	599	0.023136236	0.113055273	xyloglucan-specific endo-beta-1,4-glucanase
ecd3_2_1_54	0.000465379	0.000205	311	0.023389883	0.113485486	cyclomaltodextrinase
ecd1_3_3_4	0.000709256	0.000313	578	0.023736808	0.114859543	protoporphyrinogen oxidase
ecd3_4_13_21	-0.000581494	0.000257	803	0.02380187	0.115073606	dipeptidase e
ecd4_2_1_7	0.000582399	0.000258	895	0.023968279	0.115574792	altronate dehydratase
ecd6_3_1_2	0.000601705	0.000267	895	0.024218891	0.116174991	glutamate--ammonia ligase
ecd1_10_3_10	-0.000821538	0.000365	831	0.02468751	0.117403768	ubiquinol oxidase (h(+)-transporting)
ecd5_4_99_1	-0.000814173	0.000364	529	0.025538877	0.120420446	methylaspartate mutase
ecd3_2_1_78	0.001421238	0.00064	798	0.026732902	0.12393122	mannan endo-1,4-beta-mannosidase
ecd1_7_1_7	-0.000504605	0.000228	823	0.027182113	0.125592269	gmp reductase
ecd2_4_1_83	-0.00041385	0.000188	842	0.028081858	0.128383687	dolichyl-phosphate beta-d-mannosyltransferase
ecd3_5_4_2	-0.000638665	0.000291	895	0.028252063	0.128730089	adenine deaminase
ecd3_2_1_20	0.000627507	0.000287	895	0.029123418	0.131269173	alpha-glucosidase
ecd3_5_4_28	-0.000649842	0.000297	895	0.029116867	0.131269173	s-adenosylhomocysteine deaminase
ecd6_3_4_18	-0.000682081	0.000315	894	0.030846522	0.137456025	5-(carboxyamino)imidazole ribonucleotide synthase
ecd2_2_1_2	-0.000757022	0.00035	895	0.030998913	0.137684878	transaldolase
ecd2_6_1_62	-0.000685641	0.000318	895	0.031204972	0.138393802	adenosylmethionine--8-amino-7-oxononanoate transam
ecd4_2_1_75	-0.000639491	0.000299	753	0.032617749	0.142478094	uroporphyrinogen-iii synthase
ecd2_8_3_9	0.000393853	0.000185	802	0.033557047	0.145668688	butyrate--acetoacetate coa-transferase
ecd2_1_1_77	-0.000614657	0.000289	658	0.033855103	0.146387557	protein-l-isoaspartate(d-aspartate) o-methyltransf
ecd2_7_1_53	-0.000596498	0.000284	791	0.035946086	0.152643213	l-xylulokinase
ecd3_2_1_31	0.000851799	0.000408	895	0.037267662	0.155779954	beta-glucuronidase

ecd1_1_1_271	0.001021403	0.000494	895	0.038902868	0.159662824	gdp-l-fucose synthase
ecd1_7_7_1	0.00034679	0.000169	738	0.040282066	0.162855846	ferredoxin--nitrite reductase
ecd3_4_24_70	-0.000683111	0.000336	629	0.042063573	0.167105179	oligopeptidase a
ecd4_1_99_1	-0.000720409	0.000355	889	0.042730685	0.169023777	tryptophanase
ecd2_7_7_63	-0.00052477	0.000268	895	0.050705631	0.190054343	lipoate--protein ligase
ecd5_1_3_20	-0.000690992	0.000353	879	0.050698177	0.190054343	adp-glyceromanno-heptose 6-epimerase
ecd3_1_4_37	0.000587661	0.000301	306	0.051274307	0.190850715	2',3'-cyclic-nucleotide 3'-phosphodiesterase
ecd5_1_3_4	-0.000800633	0.000411	894	0.051519397	0.191134319	l-ribulose-5-phosphate 4-epimerase
ecd5_4_2_2	-0.000707494	0.000364	895	0.052005026	0.192098781	phosphoglucomutase (alpha-d-glucose-1,6-bisphospha
ecd1_4_3_16	-0.000683831	0.000354	895	0.053754255	0.197110826	l-aspartate oxidase
ecd3_1_6_12	0.000456797	0.000237	830	0.054156682	0.197988106	n-acetylgalactosamine-4-sulfatase
ecd4_2_2_2	-0.000398867	0.000207	564	0.054353562	0.198009362	pectate lyase
ecd3_2_1_156	0.000497168	0.000263	350	0.058993	0.208037854	oligosaccharide reducing-end xylanase
ecd3_6_3_16	-0.000498778	0.000266	796	0.061509614	0.21408857	arsenite-transporting atpase
ecd3_5_1_5	-0.00096094	0.000514	887	0.061742065	0.2145828	urease
ecd3_4_13_3	0.000480765	0.000257	892	0.062155312	0.215476948	transferred entry: 3.4.13.18 and 3.4.13.20
ecd3_5_4_9	-0.000457086	0.000245	869	0.062382313	0.216128315	methenyltetrahydrofolate cyclohydrolase
ecd2_6_1_1	0.000534957	0.000287	895	0.062662506	0.216420629	aspartate transaminase
ecd1_11_1_5	-0.000598585	0.000324	800	0.064623371	0.220965568	cytochrome-c peroxidase
ecd3_3_1_1	0.000683149	0.000369	895	0.064738193	0.220965568	adenosylhomocysteinase
ecd2_7_1_21	0.000699974	0.000379	895	0.064987868	0.220998743	thymidine kinase
ecd3_1_26_12	-0.000308783	0.000167	555	0.065586534	0.221806112	ribonuclease e
ecd4_1_1_50	0.000685801	0.000372	886	0.065662136	0.221816372	adenosylmethionine decarboxylase
ecd4_2_1_59	-0.000636738	0.000347	895	0.066823079	0.223932282	3-hydroxyacyl-[acyl-carrier-protein] dehydratase
ecd2_1_1_14	0.000559672	0.000305	893	0.066992561	0.224364178	5-methyltetrahydropteroyltriglutamate--homocystein
ecd4_2_99_20	-0.000399461	0.000218	504	0.067068729	0.224483219	2-succinyl-6-hydroxy-2,4-cyclohexadiene-1-carboxyl
ecd2_7_7_58	-0.000678631	0.000371	753	0.067755714	0.225825136	(2,3-dihydroxybenzoyl)adenylate synthase
ecd3_6_1_22	0.000495261	0.000271	895	0.067996574	0.2264913	nad(+) diphosphatase
ecd5_1_3_9	-0.000507802	0.000278	894	0.068202377	0.226494192	n-acylglucosamine-6-phosphate 2-epimerase
ecd2_3_1_47	-0.000553581	0.000303	895	0.068394353	0.226632739	8-amino-7-oxononanoate synthase

ecd2_7_1_2	0.000576834	0.000316	895	0.068521223	0.226788136	glucokinase
ecd2_7_8_6	-0.000781949	0.000429	894	0.068658727	0.227054533	undecaprenyl-phosphate galactose phosphotransferase
ecd3_5_3_11	0.000474008	0.000261	894	0.069743109	0.229132236	agmatinase
ecd3_4_11_7	-0.000583068	0.000322	663	0.070117602	0.229407857	glutamyl aminopeptidase
ecd2_4_1_187	-0.000524659	0.00029	761	0.07044963	0.230085493	mannosaminyltransferase
ecd3_5_3_12	0.000632947	0.00035	894	0.071160816	0.231437146	agmatine deiminase
ecd5_4_99_5	0.000647154	0.000359	894	0.071398872	0.231522339	chorismate mutase
ecd3_4_14_5	-0.000371525	0.000206	684	0.071932856	0.232368565	dipeptidyl-peptidase Iv
ecd1_3_3_1	0.000540662	0.000301	891	0.07247512	0.233526248	transferred entry: 1.3.98.1
ecd4_1_1_18	-0.000543244	0.000303	894	0.073530508	0.235433774	lysine decarboxylase
ecd5_4_2_6	-0.000461863	0.000258	886	0.074186641	0.23673917	beta-phosphoglucomutase
ecd2_4_1_129	-0.000621959	0.00035	895	0.076205532	0.240772882	peptidoglycan glycosyltransferase
ecd1_1_1_140	-0.000494767	0.00028	812	0.077091182	0.242791824	sorbitol-6-phosphate 2-dehydrogenase
ecd6_4_1_2	0.000725197	0.000412	895	0.078887477	0.245944162	acetyl-coa carboxylase
ecd3_5_1_11	0.000389148	0.000223	892	0.080697338	0.249543084	penicillin amidase

Supplemental Table 2: List of species that were significantly associated with metabolic risk (METscore).

Species	beta coefficients	SEM	N	N.not.0	P-value	Q-value
Coprococcus_sp_art55_1	-0.01421	0.004108	911	148	0.000586	0.037377
Faecalibacterium_prausnitzii	-0.01388	0.004157	911	901	0.000919	0.044591
Acidaminococcus_unclassified	0.004625	0.001507	911	116	0.002217	0.067882
Eubacterium_ventriosum	0.004583	0.001522	911	530	0.002728	0.080408
Subdoligranulum_unclassified	-0.01346	0.00461	911	910	0.003648	0.098012
Butyrivibrio_crossotus	-0.01102	0.003807	911	121	0.00392	0.098339
Barnesiella_intestinihominis	-0.00868	0.003272	911	482	0.008141	0.142258
Bacteroides_xylanisolvens	0.0041	0.001561	911	639	0.008861	0.14569
Clostridium_bolteae	0.004291	0.001786	911	454	0.016651	0.211992
Eubacterium_rectale	0.017374	0.007155	911	868	0.015593	0.211992
Paraprevotella_unclassified	0.004809	0.002	911	255	0.016409	0.211992
Roseburia_inulinivorans	0.005544	0.002337	911	761	0.018104	0.218347
Streptococcus_vestibularis	0.001555	0.000665	911	137	0.019922	0.230273
Burkholderiales_bacterium_1_1_47	0.002128	0.000932	911	406	0.022708	0.236928

Analyzed using MaAsLin2 adjusted for repeated measurements (participants ID as random intercept), age, energy intake, alcohol, smoking, physical activity, using of antibiotics, consumed any probiotics and fecal sample characteristics)