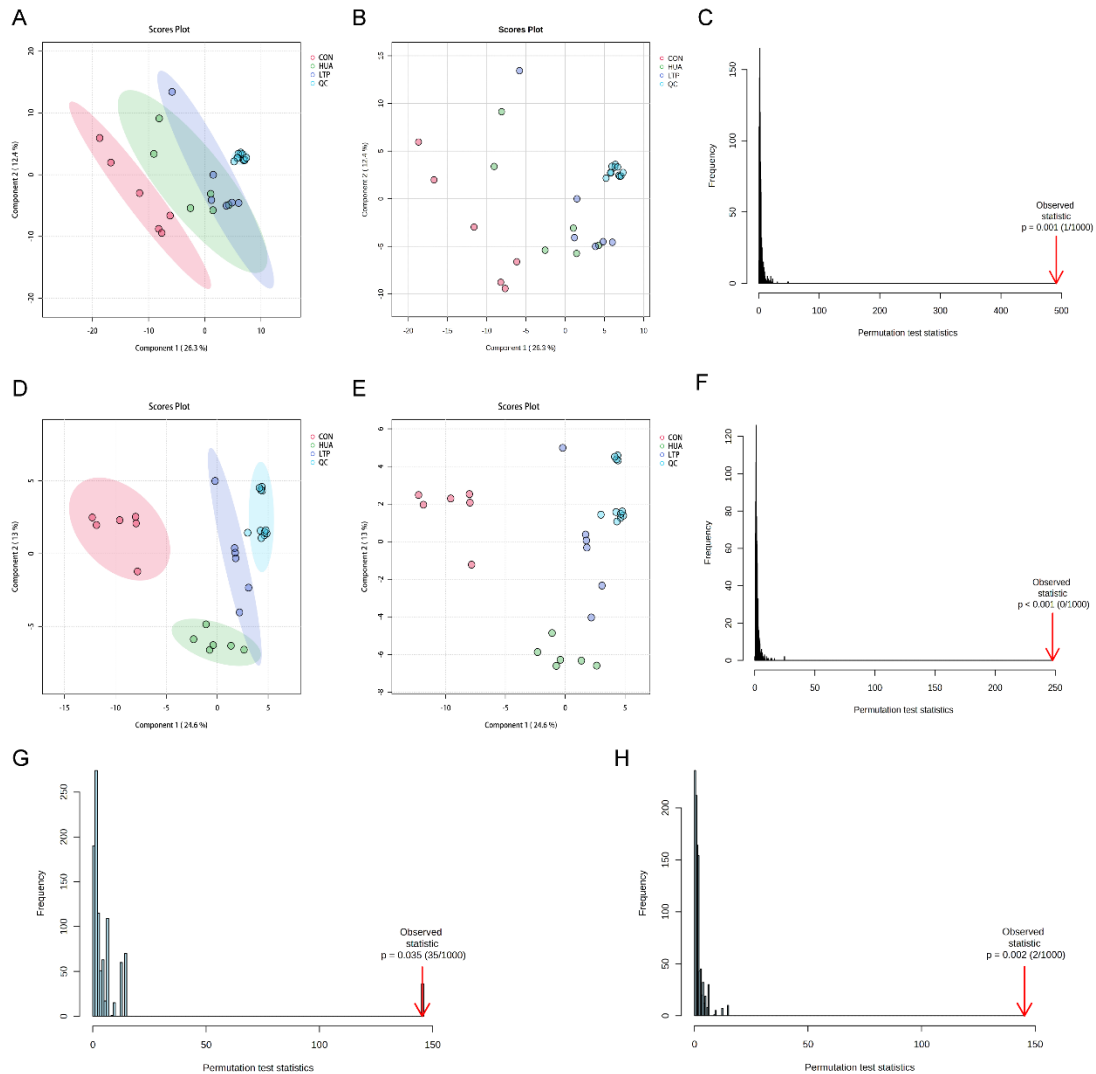
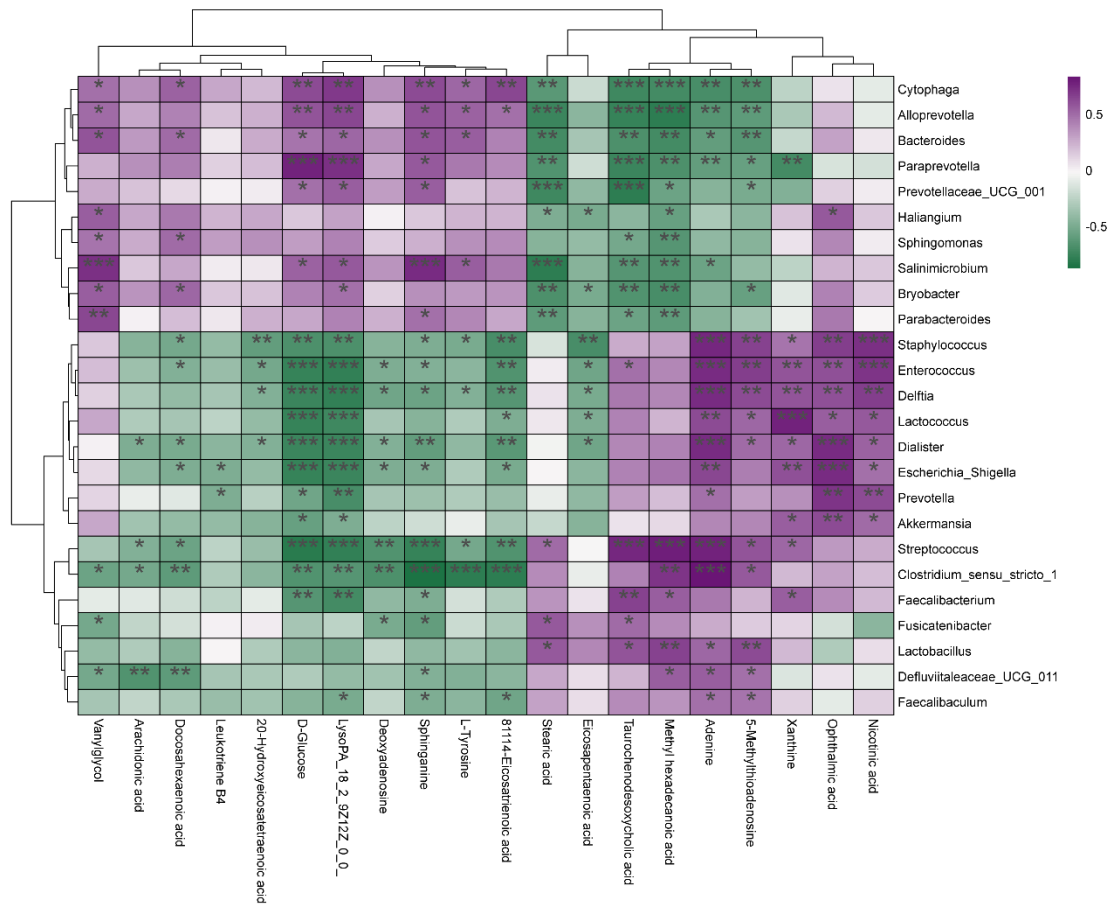


Supplementary Figure 1 Alpha rarefaction Curve, ROC curve for PLS-DA, permanova statistics for PCoA plots. (A,B) Alpha rarefaction Curve of all samples. (C) ROC curve for PLS-DA. (D-G) permanova statistics for PCoA plots among all group.



Supplementary Figure 2 Data assessment of Con, HUA and LTP group samples in serum metabolite composition. (A-B and D-E) PLS-DA score plot of all analyzed samples (including the quality control samples; A-B. ESI+, D-E. ESI-). (C,F) Permutation test charts for partial least squares discriminant analysis model of all analyzed samples (C. ESI+, F. ESI-, $n=1000$). (G,H) Permutation test charts for partial least squares discriminant analysis model of CON, HUA, and LTP group (G. ESI+, H. ESI-, $n=1000$).



Supplementary Figure 3 Correlation analysis between gut microbiota and metabolites from all group. The R values are represented by gradient colors, where purple and green cells indicate positive and negative correlations, respectively; * $p < 0.05$, ** $p < 0.01$, *** $p < 0.001$.

Supplementary Table 3 Differential Metabolic Biomarkers in Hyperuricemic Mice Treated with *Poecilobdella manillensis* Total Protein Extract

Differential Metabolic Biomarkers in Hyperuricemic Mice Treated with <i>Poecilobdella manillensis</i> Total Protein Extract										
NO	identification	formula	CON vs HUA			raw.pval	HUA vs LTP			
			VIP	FC	trend		VIP	FC	trend	raw.pval
1	Deoxyadenosine	C10 H13 N5 O3	1.08901971	0.64845	↓	0.093074	1.39242482	2.7155	↑ *	0.034026
2	5'-Methylthioadenosine	C11 H15 N5 O3 S	1.4792507	1.5364	↑ *	0.041126	1.90719086	0.29651	↓ **	0.005831
3	Adenine	C5 H5 N5	1.7031755	2.4785	↑ **	0.008658	2.18771144	0.19589	↓ ***	4.91E-05
4	Arachidonic acid	C20 H32 O2	1.58000534	0.45268	↓ *	0.025974	1.05915863	2.5691	↑ **	0.002182
5	Docosahexaenoic acid	C23 H34 O2	1.40930888	0.41083	↓ *	0.015152	1.30737335	2.7763	↑	0.076189
6	Nicotinic acid	C6 H5 N O2	1.21966329	0.32188	↓ *	0.041126	1.604527	0.30586	↓ *	0.042866
7	ophthalmic acid	C11 H19 N3 O6	1.43710613	0.52005	↓ *	0.025974	1.45715114	0.48186	↓ *	0.034013
8	Methyl hexadecanoic acid	C16 H32 O2	1.01730576	0.57625	↓ *	0.015152	1.50464373	2.6738	↑ *	0.041463
9	20-Hydroxyeicosatetraenoic a	C20 H32 O3	1.28286573	0.61554	↓ *	0.015152	1.69435078	2.3667	↑ **	0.008804
10	8,11,14-Eicosatrienoic acid	C20 H34 O3	1.53141706	0.4396	↓ *	0.015152	1.82707704	2.2423	↑ **	0.003609
11	5-Hydroxyindoleacetic acid	C10 H9 N O3	1.15064324	1.8549	↑ *	0.041126	1.71309667	0.1804	↓ **	0.007821
12	Eicosapentaenoic acid	C20 H30 O2	1.44979605	2.4731	↑ *	0.015152	1.29602732	1.6579	↑	0.074653
13	Leukotriene B4	C20 H32 O4	1.06809404	0.57877	↓	0.064935	1.57653439	2.5789	↑ *	0.01944
14	LysoPA(18:2(9Z,12Z)/0:0)	C21H41O7P	1.81427829	0.034336	↓ **	0.002165	2.24991255	89.942	↑ ***	3.59E-07
15	5a-Pregnane-3,20-dione	C21 H32 O2	1.67635594	3.1912	↑ **	0.002165	1.39923672	1.8446	↑ *	0.048066
16	Stearic acid	C18 H36 O2	1.56968283	2.1151	↑ **	0.002165	2.00454318	0.554	↓ ***	0.000339
17	Taurochenodesoxycholic acid	C26 H45 N O6 S	1.79495908	36.771	↑ **	0.002165	2.23529798	0.035891	↓ ***	1.19E-06
18	Vanylglycol	C9H12O4	1.95651094	0.001764	↓ **	0.002165	1.23801871	1.5648	↑	0.071106
19	Xanthine	C5 H4 N4 O2	1.42812841	0.62297	↓ **	0.008658	1.15912088	0.005904	↓ *	0.013658
20	sphinganine	C18H39NO2	2.18774056	0.007657	↓ **	0.002165	2.02935705	13.067	↑ ***	0.000159
21	D-Glucose	C6 H12 O6	1.74262168	0.16137	↓ **	0.002165	1.77068765	2.8678	↑ **	0.00507
22	L-Tyrosine	C9 H11 N O3	1.56604439	0.31993	↓	0.064935	1.63727876	3.9373	↑ *	0.01229

Supplementary Table 4 P value matrix for correlation analysis between gut microbiota and metabolites from all groups

	Adenine	Deoxyade	D- Glucose	L- Tyrosine	Sphingani	Xanthine	Stearic aci	Taurocher	Arachidon	5- Methylt	Vanylglyc	Eicosapen	Docosahe	Ophthalm	LysoPA_18	Methyl he	Leukotrier	20-Hydro	81114-Eic	Nicotinic ε
Bryobacte	0.052224	0.602304	0.086179	0.180485	0.129067	0.685431	0.002026	0.003567	0.16156	0.01349	0.016546	0.030715	0.028312	0.086179	0.041414	0.001642	0.506785	0.457929	0.145334	0.549297
Bacteroid	0.011451	0.127873	0.048169	0.016038	0.011451	0.413635	0.001037	0.00236	0.168071	0.004322	0.010743	0.170838	0.03614	0.229322	0.025867	0.001089	0.886853	0.287484	0.077783	0.867607
Alloprevot	0.004002	0.315513	0.009747	0.020783	0.010403	0.200205	0.000335	0.000233	0.261026	0.008254	0.031069	0.065782	0.081019	0.376225	0.004663	7.30E-05	0.488839	0.307345	0.042836	0.729259
Paraprevot	0.009128	0.252992	0.000134	0.057548	0.012803	0.001075	0.003202	0.000338	0.139849	0.013621	0.036049	0.476122	0.071463	0.551338	0.000716	0.002342	0.605185	0.448039	0.121276	0.502387
Prevotella	0.043367	0.187119	0.019899	0.224953	0.14364	0.127023	0.759685	0.190098	0.765921	0.190098	0.63835	0.087065	0.644243	0.001001	0.001685	0.417463	0.03844	0.27492	0.107706	0.006567
Prevotella	0.055216	0.197128	0.042836	0.478445	0.018028	0.052786	0.000622	7.30E-05	0.483628	0.035254	0.303309	0.081019	0.698742	0.598461	0.017513	0.015569	0.983788	0.912608	0.367178	0.912608
Parabacte	0.06301	0.33649	0.246615	0.102535	0.037965	0.778883	0.006711	0.017513	0.977304	0.162629	0.004322	0.059017	0.423288	0.060326	0.087785	0.006711	0.874015	0.328004	0.253756	0.996757
Cytophag	0.001499	0.131143	0.005	0.024839	0.00524	0.307781	0.005489	0.000623	0.125364	0.001885	0.042839	0.439052	0.023189	0.824831	0.001781	0.000865	0.268362	0.402314	0.004769	0.72213
Salinimicro	0.018755	0.163111	0.023254	0.018462	0.000372	0.318564	9.34E-05	0.004492	0.5148	0.093925	0.000643	0.059647	0.243016	0.3394	0.012878	0.003046	0.900559	0.868825	0.060369	0.538564
Faecalibac	0.039419	0.348671	0.058654	0.071319	0.042402	0.606176	0.237963	0.110635	0.366561	0.045555	0.166114	0.764009	0.192211	0.714154	0.030623	0.16066	0.119173	0.14756	0.020952	0.617846
Enterococ	0.000498	0.041449	0.000239	0.076077	0.047404	0.008835	0.90798	0.04033	0.130989	0.001824	0.435644	0.019748	0.04996	0.007874	0.000122	0.101966	0.113435	0.026162	0.003559	0.000856
Lactobacil	0.024514	0.349454	0.074643	0.162629	0.079389	0.390027	0.012582	0.010743	0.232717	0.005215	0.106485	0.091318	0.061657	0.236143	0.052786	0.002791	0.964342	0.22596	0.076201	0.766392
Lactococ	0.006295	0.169753	0.000206	0.223716	0.058919	0.000161	0.873269	0.087625	0.232234	0.025698	0.268362	0.03791	0.169753	0.016182	0.000623	0.344956	0.307781	0.09441	0.03791	0.013374
Streptoco	0.000158	0.003248	4.40E-05	0.027745	0.000267	0.02556	0.036092	0.000285	0.047369	0.010989	0.171427	0.923274	0.016567	0.16857	0.000169	9.63E-05	0.330286	0.120305	0.003384	0.304756
Staphyloc	0.000237	0.054923	0.002123	0.036582	0.040501	0.048619	0.555849	0.293938	0.056403	0.002872	0.50415	0.001271	0.024126	0.001913	0.001964	0.234655	0.100494	0.00976	0.001421	0.000795
Clostridiu	1.50E-05	0.002111	0.002629	8.94E-05	2.59E-06	0.404216	0.113691	0.086217	0.02556	0.012118	0.017594	0.76594	0.008384	0.221619	0.004482	0.001088	0.228496	0.068298	7.69E-05	0.439152
Defluvital	0.017319	0.146658	0.227149	0.050287	0.048551	0.597438	0.251904	0.340319	0.003525	0.03717	0.024074	0.774877	0.008451	0.822981	0.102518	0.015594	0.176627	0.213732	0.074171	0.687256
Fusicateni	0.296293	0.03055	0.166785	0.448877	0.010146	0.672764	0.012266	0.031498	0.37218	0.555849	0.026138	0.139169	0.532946	0.532946	0.325437	0.116164	0.953676	0.899819	0.209067	0.072903
Faecalibac	0.05868	0.088473	0.004291	0.52971	0.039922	0.016669	0.146497	0.002601	0.644474	0.348359	0.690663	0.834998	0.491595	0.118721	0.001134	0.018178	0.317999	0.694262	0.232619	0.380317
Dialister	0.000411	0.04996	0.000467	0.087343	0.009888	0.027785	0.878714	0.10419	0.04033	0.036083	0.988936	0.020392	0.030367	0.000438	0.000467	0.081567	0.065918	0.048669	0.005485	0.023866
Haliangiur	0.199223	0.9593	0.506785	0.356206	0.526428	0.495719	0.040384	0.072653	0.257183	0.076699	0.01813	0.032402	0.062873	0.013707	0.242257	0.013062	0.363176	0.268747	0.344759	0.526428
Sphingom	0.0859	0.543663	0.189185	0.132838	0.33559	0.795637	0.057358	0.026838	0.294086	0.059971	0.046644	0.106533	0.031427	0.096757	0.089415	0.003665	0.198359	0.142533	0.112727	0.924739
Delftia	0.000385	0.023866	0.000337	0.046164	0.017329	0.008835	0.799106	0.058253	0.208962	0.005044	0.59981	0.03313	0.091354	0.006999	0.000132	0.095498	0.187988	0.048669	0.003891	0.001918
Escherichi	0.004725	0.042906	0.00031	0.221318	0.044007	0.007223	0.954987	0.089991	0.088111	0.072458	0.718006	0.075744	0.037712	0.000351	0.000449	0.052328	0.036733	0.099828	0.031267	0.03871
Akkerman	0.101761	0.314479	0.013983	0.753462	0.482709	0.018263	0.384388	0.828986	0.153782	0.092382	0.267395	0.061055	0.101761	0.005095	0.041342	0.734881	0.101761	0.063769	0.187119	0.035697

Supplementary Table 5 Spearman rank correlation matrix for Correlation analysis between gut microbiota and metabolites from all groups

	Adenine	Deoxyade	D-Glucose	L-Tyrosine	Sphingani	Xanthine	Stearic aci	Taurocher	Arachidon	5-Methylt	Vanylglyc	Eicosapen	Docosahe	Ophthalm	LysoPA 1&	Methyl he	Leukotrier	20-Hydro	81114-Eic	Nicotinic a
Bryobacte	-0.46434	0.131743	0.415745	0.330436	0.371471	-0.10259	-0.67707	-0.64899	0.344475	-0.57016	0.556127	-0.50969	0.516172	0.415745	0.484856	-0.68679	0.167378	0.186815	0.357433	0.151118
Bacteroid	-0.58101	0.372549	0.47162	0.558308	0.581011	-0.20537	-0.70691	-0.66976	0.339525	-0.6388	0.585139	-0.33746	0.496388	0.298246	0.52322	-0.70485	0.03612	0.265222	0.426213	0.042312
Alloprevot	-0.64293	0.250774	0.591331	0.539732	0.587203	-0.31682	-0.75026	-0.76264	0.27967	-0.60165	0.508772	-0.44272	0.422085	0.221878	0.634675	-0.79773	0.174407	0.254902	0.48194	-0.08772
Paraprevo	-0.59544	0.284236	0.780093	0.4554	0.573659	-0.7054	-0.65457	-0.75001	0.362038	-0.56951	0.240667	-0.17946	0.434653	-0.15042	0.722001	-0.67013	0.130707	0.190874	0.378636	-0.16909
Prevotella	0.48087	-0.32575	-0.54292	-0.30093	-0.35884	0.37332	-0.07756	0.323682	-0.07549	0.323682	0.118925	-0.41469	-0.11686	0.708378	-0.68563	0.203723	-0.49121	-0.27198	-0.39193	0.615306
Prevotella	-0.45924	0.318885	0.48194	0.178535	0.550052	-0.46336	-0.72755	-0.79773	0.176471	-0.49845	0.256966	-0.42208	0.098039	0.133127	0.552116	-0.56037	0.00516	0.027864	0.226006	0.027864
Parabacte	-0.44685	0.240454	0.287926	0.397317	0.49226	-0.07121	-0.61404	-0.55212	0.007224	-0.34365	0.638803	-0.45304	0.201238	0.45098	0.413829	-0.61404	0.040248	0.244582	0.283798	-0.00103
Cytophag	-0.6909	0.369613	0.630823	0.526339	0.628211	-0.25468	-0.6256	-0.72747	0.374837	-0.68045	0.481933	-0.1946	0.531563	0.05616	0.683065	-0.71441	0.275577	0.210274	0.633435	-0.09012
Salinimicr	-0.54721	0.343283	0.531353	0.548347	0.746613	-0.24925	-0.7908	-0.63672	0.164278	-0.40673	0.72622	-0.45205	0.290035	0.239052	0.573272	-0.65711	0.031723	0.041919	0.450913	0.155214
Faecalibac	0.489078	-0.23463	-0.45362	-0.43485	-0.48282	0.130351	0.293029	0.388968	-0.22629	0.476564	-0.341	0.076125	-0.32223	-0.09281	-0.50993	0.34517	-0.38063	-0.3556	-0.53913	0.12618
Enterococ	0.735978	-0.48478	-0.7618	-0.42844	-0.47304	0.597468	0.029345	0.48713	-0.36975	0.681983	0.196026	-0.54347	-0.46835	0.604511	-0.78293	0.397921	-0.38618	-0.52234	-0.64912	0.714849
Lactobacil	0.527348	-0.23426	-0.43034	-0.34365	-0.42415	0.215686	0.574819	0.585139	-0.29618	0.628483	-0.39319	0.409701	-0.44892	-0.29412	-0.46336	0.661507	-0.01135	-0.30031	-0.42828	0.075335
Lactococc	0.617763	-0.33827	-0.76665	-0.3017	-0.4532	0.774489	0.040488	0.414018	-0.29647	0.523727	0.275577	-0.49238	-0.33827	0.557684	-0.72747	0.236395	-0.25468	-0.40618	-0.49238	0.570745
Streptococ	0.775051	-0.65385	-0.8112	-0.51776	-0.75804	0.524143	0.4965	0.755914	-0.47311	0.58368	-0.33702	-0.02445	-0.55604	0.339151	-0.77292	0.789935	-0.24347	-0.37955	-0.65172	0.256224
Staphyloc	0.762113	-0.45973	-0.67484	-0.49537	-0.48677	0.470789	-0.14873	0.261823	-0.45727	0.660088	0.168402	-0.69819	-0.52856	0.679756	-0.67853	0.295012	-0.39949	-0.59125	-0.69328	0.717861
Clostridiu	0.836715	-0.67511	-0.66448	-0.79206	-0.87074	0.209444	0.385931	0.415699	-0.52414	0.577301	-0.55179	-0.07549	-0.60069	0.303003	-0.63684	0.704882	-0.29875	-0.43909	-0.79631	0.19456
Defluviita	0.552905	-0.35634	-0.29958	-0.46776	-0.47092	-0.1335	0.284862	0.238611	-0.64961	0.49404	-0.52873	0.072529	-0.60021	0.056762	-0.39733	0.560263	-0.33321	-0.30799	-0.43097	-0.10196
Fusicateni	0.260594	-0.51012	-0.34049	-0.19053	-0.58879	0.106942	0.576502	0.507666	-0.22372	0.148735	-0.52242	0.362618	-0.15734	-0.15734	-0.24584	0.383515	0.014751	0.03196	-0.31099	-0.43268
Faecalibac	0.45358	-0.41302	-0.63919	-0.15857	-0.488	0.555605	0.356472	0.665005	-0.11678	0.23478	-0.1008	0.052856	-0.17332	0.381057	-0.70311	0.549459	-0.24953	-0.09957	-0.29624	0.220029
Dialister	0.743021	-0.46835	-0.73833	-0.41435	-0.59043	0.517649	-0.03874	0.395573	-0.48713	0.496521	0.003521	-0.54113	-0.51061	0.740673	-0.73833	0.421397	-0.44253	-0.4707	-0.62564	0.529388
Haliangiur	-0.31748	0.012958	0.167378	0.23109	0.159819	0.171697	-0.48702	-0.43302	0.281843	-0.42762	0.549648	-0.50537	0.447061	0.569085	0.290482	-0.57232	0.22785	0.275364	0.236489	0.159819
Sphingom	-0.41608	0.153293	0.324314	0.368112	0.240889	0.065697	-0.45571	-0.52036	0.261745	-0.45154	0.474478	-0.39314	0.507848	0.403567	0.41191	-0.64758	0.318057	0.359769	0.386882	0.023985
Delftia	0.745368	-0.52939	-0.75006	-0.47539	-0.55286	0.597468	0.064559	0.454264	-0.31106	0.630335	0.13264	-0.50356	-0.40966	0.611554	-0.78058	0.404964	-0.32514	-0.4707	-0.64442	0.679635
Escherichi	0.633946	-0.4818	-0.75302	-0.30319	-0.47959	0.60969	-0.01433	0.411238	-0.41344	0.433288	0.091509	-0.42888	-0.49282	0.748607	-0.73979	0.464159	-0.49503	-0.40021	-0.50826	0.490619
Akkerman	0.398139	-0.25129	-0.56774	-0.07963	-0.17684	0.549122	-0.2182	0.054809	-0.35057	0.408481	0.276112	-0.44985	-0.39814	0.629784	-0.48501	0.085833	-0.39814	-0.44571	-0.32575	0.497416

Supplementary Table 6 P value matrix for correlation analysis between the gut microbiota and metabolites from HUA and LTP groups

	Adenine	Deoxyade	D-Glucose	L-Tyrosine	Sphingani	Xanthine	Stearic aci	Taurocher	Arachidon	5-Methylt	Vanylglyc	Eicosapen	Docosahe	Ophthalm	LysoPA_1&	Methyl he	Leukotrier	20-Hydro	81114-Eic	Nicotinic e
Bryobacte	0.03152	0.335508	0.077052	0.150935	0.202808	0.126919	0.105548	0.004117	0.185824	0.010054	0.871178	0.648215	0.084229	0.624676	0.010054	0.03152	0.225327	0.173686	0.041075	0.074756
Bifidobact	0.024003	0.137658	0.022034	0.137658	0.03317	0.151735	0.275567	0.041521	0.1667	0.070894	0.075362	0.378569	0.044609	0.124455	0.010245	0.112109	0.391097	0.245265	0.026097	0.275567
Collinsella	0.025161	0.016989	0.008624	0.17281	0.012286	0.263403	0.511289	0.027585	0.06182	0.190954	0.125051	0.210231	0.057492	0.003803	0.003803	0.012286	0.01373	0.125051	0.025161	0.376781
Bacteroid	0.00026	0.013906	0.00824	0.00026	0.002586	0.124455	0.1667	0.044609	0.015317	0.024003	0.084869	0.070894	0.003446	0.226206	0.000332	0.00824	0.118176	0.011374	0.002586	0.030676
Alloprevot	0.001615	0.151735	0.018453	0.044609	0.024003	0.03317	0.013906	0.000521	0.118176	0.002993	0.556737	0.617523	0.084869	0.354148	0.0019	0.004508	0.378569	0.137658	0.013906	0.016831
Paraprevot	0.127893	0.141425	0.010567	0.251553	0.052774	0.000158	0.004078	0.011731	0.082308	0.068541	0.69463	0.810876	0.134549	0.213512	0.022709	0.072939	0.844854	0.662387	0.45353	0.362722
Prevotella	0.000521	0.006543	0.0019	0.084869	0.000521	0.151735	0.208063	0.016831	0.013906	0.054842	0.041521	0.21702	0.02832	0.000332	0.001615	0.007355	0.118176	0.095157	0.030676	0.199335
Prevotella	0.084869	0.245265	0.058589	0.571701	0.030676	0.002223	0.001363	0.000643	0.319139	0.112109	0.648796	0.648796	0.680598	0.208063	0.038588	0.041521	0.845931	0.948402	0.470532	0.330589
Rikenellac	0.230003	0.086718	0.059984	0.150935	0.290588	0.274671	0.556033	0.290588	0.015807	0.054324	0.795063	0.211643	0.002512	0.126919	0.185824	0.047374	0.143798	0.254246	0.120547	0.244376
Parabacte	0.022034	0.226206	0.354148	0.022034	0.080019	0.633091	0.255138	0.391097	0.96559	0.527302	0.030676	0.862898	0.648796	0.931234	0.054842	0.144586	0.416775	0.182564	0.054842	0.190836
Bilophila	0.001481	0.072334	0.105589	0.027655	0.026537	0.480487	0.144147	0.170335	0.208138	0.047092	0.039299	0.606586	0.147719	0.133768	0.089184	0.012128	0.494665	0.178331	0.016296	0.03509
Faecalibac	0.085853	0.266468	0.071528	0.073792	0.062948	0.217296	0.732825	0.266468	0.166105	0.174049	0.293268	0.339168	0.065022	0.549541	0.047978	0.076106	0.261285	0.053261	0.028117	0.051457
Enterococ	0.001167	0.027866	0.00096	0.003736	0.004947	0.035814	0.058366	0.001998	0.002172	0.000448	0.447411	0.040343	0.001683	0.089297	0.002358	0.001167	0.144257	0.033005	0.00096	0.004015
Lactobacil	0.095157	0.745609	0.208063	0.330589	0.265239	0.235621	0.416775	0.066612	0.190836	0.013906	0.982792	0.712884	0.124455	0.845931	0.151735	0.095157	0.778725	0.182564	0.075362	0.016831
Lactococc	0.025161	0.111294	0.000916	0.035863	0.012286	0.000463	0.006684	0.002354	0.081393	0.018816	0.699238	0.363094	0.053379	0.24128	0.00198	0.022896	0.434119	0.230645	0.035863	0.06182
Clostridia	0.084869	0.080019	0.010245	0.062511	0.016831	0.005801	0.007355	0.051266	0.391097	0.378569	0.296904	0.512841	0.330589	0.226206	0.007355	0.208063	0.586824	0.586824	0.226206	0.617523
Clostridiu	0.000916	0.00198	0.057492	0.001654	0.002354	0.495392	0.230645	0.16416	0.057492	0.071145	0.009739	0.125051	0.025161	0.04578	0.042283	0.003803	0.274889	0.132328	0.00137	0.081393
Faecalibac	0.04689	0.027344	0.010093	0.296896	0.005872	0.015352	0.030094	0.003707	0.271611	0.358405	0.426064	0.887689	0.380293	0.027344	0.000988	0.028695	0.23046	0.596082	0.23046	0.777247
Dialister	0.00096	0.001683	0.001167	0.015905	0.000179	0.113004	0.097752	0.004309	0.022287	0.040343	0.064714	0.071526	0.025526	0.001683	0.001683	0.001407	0.081352	0.126281	0.001407	0.116233
Haliangiur	0.263447	0.922389	0.974099	0.300583	0.845395	0.428244	0.408574	0.616271	0.284326	0.201942	0.948221	0.334615	0.129422	0.370632	0.279021	0.421637	0.078773	0.021417	0.23385	0.180877
Sphingom	0.359293	0.93971	0.688104	0.274671	0.93971	0.753434	0.390234	0.736944	0.323944	0.285226	0.786694	0.365374	0.089257	0.435736	0.198476	0.490702	0.054324	0.018175	0.216147	0.207197
Delftia	0.000425	0.00757	0.001015	0.001231	0.000291	0.014089	0.008044	0.003626	0.024036	0.006685	0.192878	0.150973	0.014089	0.073217	0.001231	0.001119	0.338525	0.128886	0.001769	0.024036
Escherichi	0.016163	0.004017	6.92E-05	0.067587	0.002591	0.023429	0.12857	0.008563	0.01955	0.081657	0.257052	0.236742	0.016163	0.001882	0.000386	0.001589	0.023429	0.142564	0.023429	0.246774
Akkerman	0.118176	0.174519	0.000201	0.182564	0.095157	0.020185	0.265239	0.106252	0.02832	0.054842	0.571701	0.319139	0.015317	0.03317	0.026097	0.058589	0.080019	0.112109	0.124455	0.118176

Supplementary Table 7 Spearman rank correlation matrix for correlation analysis between the gut microbiota and metabolites from HUA and LTP groups

	Adenine	Deoxyade	D-Glucose	L-Tyrosine	Sphingani	Xanthine	Stearic aci	Taurocher	Arachidon	5-Methylt	Vanylglycc	Eicosapen	Docosahe	Ophthalm	LysoPA_18	Methyl he	Leukotrier	20-Hydro	81114-Eic	Nicotinic e
Bryobacte	-0.61997	0.304729	0.528897	0.441332	0.395797	-0.46585	-0.49037	-0.76007	0.409808	-0.70753	0.052539	0.147111	0.51839	-0.15762	0.707532	-0.61997	0.378284	0.420316	0.595447	-0.5324
Bifidobact	0.643357	-0.45455	-0.65035	-0.45455	-0.61538	0.440559	0.342657	0.594406	-0.42657	0.538462	-0.53147	-0.27972	-0.58741	0.468531	-0.70629	0.482517	-0.27273	-0.36364	-0.63636	0.342657
Collinsella	0.639436	-0.67063	-0.71742	-0.42109	-0.69402	0.35091	0.210546	0.631638	-0.55366	0.405496	-0.46788	-0.3899	-0.56146	0.764204	-0.7642	0.694022	-0.68622	-0.46788	-0.63944	0.280728
Bacteroides	-0.86713	0.685315	0.72028	0.867133	0.783217	-0.46853	-0.42657	-0.58741	0.678322	-0.64336	0.517483	0.538462	0.769231	-0.37762	0.86014	-0.72028	0.475524	0.699301	0.783217	-0.62238
Alloprevot	-0.8042	0.440559	0.664336	0.587413	0.643357	-0.61538	-0.68531	-0.84615	0.475524	-0.77622	0.188811	0.160839	0.517483	-0.29371	0.797203	-0.75524	0.27972	0.454545	0.685315	-0.67133
Paraprevot	-0.4648	0.450715	0.704243	0.359164	0.570437	-0.8803	-0.76058	-0.6972	0.52114	-0.54227	-0.12676	0.077467	0.457758	-0.38733	0.647903	-0.53522	0.063382	0.140849	0.239443	-0.28874
Prevotella	0.846154	-0.73427	-0.7972	-0.51748	-0.84615	0.440559	0.391608	0.671329	-0.68531	0.566434	-0.59441	-0.38462	-0.62937	0.86014	-0.8042	0.727273	-0.47552	-0.5035	-0.62238	0.398601
Prevotella	-0.51748	0.363636	0.559441	0.181818	0.622378	-0.79021	-0.81119	-0.83916	0.314685	-0.48252	-0.14685	-0.14685	0.132867	-0.39161	0.601399	-0.59441	-0.06294	0.020979	0.230769	-0.30769
Rikenellac	-0.37478	0.514887	0.556919	0.441332	0.33275	-0.34326	-0.18914	-0.33275	0.676008	-0.56743	0.084063	0.388792	0.78459	-0.46585	0.409808	-0.58144	0.448337	0.357268	0.472855	-0.36427
Parabacte	-0.65035	0.377622	0.293706	0.65035	0.524476	-0.15385	-0.35664	-0.27273	0.013986	-0.2028	0.622378	0.055944	0.146853	-0.02797	0.566434	-0.44755	0.258741	0.412587	0.566434	-0.40559
Bilophila	-0.80779	0.536176	0.490319	0.631418	0.634945	-0.22576	-0.44799	-0.4233	0.39155	-0.58203	0.599671	0.165791	0.444462	-0.45857	0.511484	-0.69491	0.218703	0.416242	0.673748	-0.61025
Faecalibac	0.516096	-0.34881	-0.53745	-0.53389	-0.55169	0.384402	0.110338	0.34881	-0.42711	0.419995	-0.33101	-0.30254	-0.54813	0.192201	-0.58016	0.530333	-0.35237	-0.56949	-0.62999	0.573044
Enterococ	0.81738	-0.63076	-0.82485	-0.76513	-0.7502	0.60837	0.55985	0.794986	-0.79125	0.850971	-0.2426	-0.59717	-0.80245	0.511329	-0.78752	0.81738	-0.44788	-0.61583	-0.82485	0.761395
Lactobacil	0.503497	-0.1049	-0.39161	-0.30769	-0.34965	0.370629	0.258741	0.545455	-0.40559	0.685315	-0.00699	-0.11888	-0.46853	0.062937	-0.44056	0.503497	-0.09091	-0.41259	-0.53147	0.671329
Lactococc	0.639436	-0.48348	-0.82659	-0.60824	-0.69402	0.849982	0.733012	0.787598	-0.52247	0.66283	-0.12477	-0.28853	-0.56925	0.366506	-0.7954	0.647234	-0.24954	-0.3743	-0.60824	0.553658
Clostridia	-0.51748	0.524476	0.706294	0.552448	0.671329	-0.74126	-0.72727	-0.57343	0.272727	-0.27972	0.328671	0.20979	0.307692	-0.37762	0.727273	-0.39161	0.174825	0.174825	0.377622	-0.16084
Clostridiu	0.826588	-0.7954	-0.56146	-0.80319	-0.7876	0.218344	0.374304	0.42889	-0.56146	0.538062	-0.70962	-0.46788	-0.63944	0.58485	-0.59265	0.764204	-0.34311	-0.46008	-0.81099	0.522466
Faecalibac	0.582464	-0.63239	-0.70728	-0.32868	-0.74056	0.678155	0.624069	0.765525	-0.34532	0.291232	-0.25379	-0.04577	-0.27875	0.63239	-0.82377	0.628229	-0.37444	-0.17058	-0.37444	0.09153
Dialister	0.824845	-0.80245	-0.81738	-0.67555	-0.8771	0.481471	0.500132	0.757663	-0.64943	0.597173	-0.54865	-0.53746	-0.63823	0.802451	-0.80245	0.809916	-0.52253	-0.46654	-0.80992	0.477738
Haliangiur	-0.35088	0.031579	-0.01053	0.326318	-0.06316	0.252633	0.26316	-0.1614	0.336844	-0.39649	0.021053	0.305265	0.463161	0.284212	0.340353	-0.25614	0.526319	0.652636	0.371932	-0.41404
Sphingom	-0.29072	0.024518	0.129597	0.343258	-0.02452	0.101576	0.273205	-0.10858	0.311734	-0.33625	0.087566	0.287216	0.511384	0.248687	0.3993	-0.22067	0.567426	0.6655	0.38529	-0.39229
Delftia	0.852672	-0.72552	-0.82275	-0.81527	-0.86389	0.684381	0.721779	0.766657	-0.64324	0.732998	-0.4039	-0.44129	-0.68438	0.53479	-0.81527	0.819014	-0.30292	-0.46373	-0.80031	0.643244
Escherichi	0.674356	-0.76137	-0.89914	-0.54384	-0.78312	0.645351	0.464073	0.717863	-0.65985	0.522082	-0.35531	-0.36981	-0.67436	0.797625	-0.85563	0.804876	-0.64535	-0.44957	-0.64535	0.362557
Akkerman	0.475524	-0.41958	-0.87413	-0.41259	-0.5035	0.657343	0.34965	0.48951	-0.62937	0.566434	-0.18182	-0.31469	-0.67832	0.615385	-0.63636	0.559441	-0.52448	-0.48252	-0.46853	0.475524

Supplementary Table 8 P value matrix for correlation analysis between gut microbiota and HUA-related parameters

	SUA	UUA	SCRE	BUN	L-XOD	K-URAT1	J-URAT1	K-ABCG2	J-ABCG2	K-Occludi	J-Occludi	K-ZO-1	J-ZO-1	K-GLUT9	J-GLUT9
Bryobacte	0.004398	0.512085	0.003596	0.050765	0.006793	0.003355	0.18088	0.010601	0.104846	0.018175	0.185824	0.00291	0.034067	0.571016	0.230003
Bifidobact	0.000521	0.862898	0.002586	0.366251	0.038588	0.005801	0.335508	0.015317	0.047374	0.044609	0.130948	0.038588	0.03317	0.245265	0.038588
Collinsella	0.010957	0.181741	0.008624	0.735559	0.06182	0.002779	0.610435	0.00198	0.016725	0.015296	0.005847	0.020783	0.053379	0.049477	0.139875
Bacteroides	0.009202	0.948402	0.00824	0.484452	0.002993	0.001615	1	0.00395	0.036759	0.001143	0.035806	0.000521	0.002993	0.21702	0.02832
Alloprevot	0.002993	0.633091	0.001363	0.084869	0.001363	0.001615	0.593787	0.022034	0.015807	0.00395	0.062511	0.005801	0.00395	0.484452	0.137658
Paraprevo	0.222673	0.187418	0.33869	0.072939	0.019022	0.022709	0.501834	0.0603	0.480499	0.134549	0.0085	0.261643	0.001962	0.163399	0.481287
Prevotella	0.009202	0.680598	0.026097	0.795415	0.047856	0.002586	0.948312	0.03317	0.000498	0.003446	0.004508	0.062511	0.012593	0.084869	0.1667
Prevotella	0.265239	0.044609	0.199335	0.151735	0.038588	0.041521	0.239528	0.174519	0.086718	0.1667	0.018453	0.416775	0.02832	0.470532	0.633091
Rikenellac	0.012375	0.99138	0.054324	0.512085	0.074756	0.147339	0.129425	0.009024	0.585491	0.140313	0.126919	0.074756	0.04259	0.023677	0.169753
Parabacte	0.124455	0.541936	0.058589	0.556737	0.199335	0.182564	0.624676	0.541936	0.063977	0.137658	0.527302	0.112109	0.151735	0.829024	0.391097
Bilophila	0.003227	0.203706	0.00743	0.70226	0.089184	0.174305	0.685466	0.399523	0.002518	0.036455	0.297865	0.147719	0.047092	0.3261	0.226449
Faecalibac	0.034354	0.817421	0.001897	0.895145	0.170048	0.083341	0.71572	0.011062	0.205065	0.104884	0.110797	0.022741	0.034354	0.363483	0.005539
Enterococ	0.006433	0.694318	0.004947	0.405191	6.81E-05	0.000782	0.472171	0.00096	0.03686	6.81E-05	0.010383	0.000565	0.002555	0.180586	0.019359
Lactobacil	0.016831	0.982792	0.0019	0.416775	0.112109	0.112109	0.244376	0.051266	0.123706	0.106252	0.391097	0.044609	0.100603	0.931234	0.038588
Lactococc	0.038979	0.376781	0.030172	0.111294	0.001125	0.00137	0.894528	0.008624	0.209362	0.01373	0.006684	0.022896	0.000359	0.210231	0.076151
Clostridia	0.354148	0.456801	0.484452	0.1667	0.026097	0.013906	0.318245	0.190836	0.409428	0.100603	0.022034	0.245265	0.012593	0.226206	0.484452
Clostridiur	0.000588	0.363094	0.002354	0.753912	0.01373	0.086876	0.847006	0.053379	0.005715	0.020783	0.155788	0.038979	0.04578	0.071145	0.042283
Faecalibac	0.10852	0.021268	0.11561	0.395258	0.034583	0.00221	0.346832	0.028695	0.061449	0.080195	0.001226	0.160048	0.015352	0.083061	0.569032
Dialister	0.003736	0.529908	0.005653	0.826649	0.001683	0.001167	0.785984	0.004309	0.002904	0.00096	0.004309	0.012252	0.016722	0.04877	0.056349
Haliangiur	0.25837	0.922389	0.168924	0.753012	0.434902	0.229118	0.149335	0.126163	0.485609	0.146543	0.811533	0.005206	0.777965	0.562806	0.49701
Sphingom	0.207197	0.948312	0.114389	0.845664	0.519302	0.181721	0.116709	0.072507	0.736497	0.189985	0.90537	0.006402	0.462837	0.845664	0.301483
Delftia	0.002675	0.880732	0.003899	0.202037	2.82E-05	0.002097	0.953924	0.006685	0.024806	0.001769	0.010753	0.008539	0.000478	0.096976	0.031225
Escherichi	0.00597	0.142564	0.008563	0.770747	0.010708	0.000486	0.761743	0.000233	0.054765	0.008563	3.27E-05	0.017798	0.000915	0.00597	0.121922
Akkerman	0.070894	0.617523	0.106252	0.96559	0.151735	0.022034	0.307014	0.044609	0.37151	0.03317	0.002993	0.112109	0.001143	0.062511	0.208063

Supplementary Table 9 Spearman rank correlation matrix for correlation analysis between gut microbiota and HUA-related parameters

	SUA	UUA	SCRE	BUN	L-XOD	K-URAT1	J-URAT1	K-ABCG2	J-ABCG2	K-Occludi	J-Occludi	K-ZO-1	J-ZO-1	K-GLUT9	J-GLUT9
Bryobacte	0.756569	-0.21016	0.767076	-0.57443	0.73205	0.770579	0.414035	-0.70403	-0.49123	-0.6655	-0.40981	-0.77758	-0.61296	0.182137	0.374782
Bifidobact	-0.84615	-0.05594	-0.78322	0.286713	-0.6014	-0.74126	-0.30473	0.678322	0.581437	0.587413	0.461538	0.601399	0.615385	-0.36364	-0.6014
Collinsella	-0.70182	0.413294	-0.71742	-0.10917	-0.55366	-0.7798	-0.16404	0.795396	0.671803	0.678426	0.74081	0.655032	0.569254	-0.57705	-0.45228
Bacteroides	0.713287	0.020979	0.72028	-0.22378	0.776224	0.804196	0	-0.76224	-0.60596	-0.81818	-0.60839	-0.84615	-0.77622	0.384615	0.629371
Alloprevot	0.776224	-0.15385	0.811189	-0.51748	0.811189	0.804196	0.171629	-0.65035	-0.67601	-0.76224	-0.55245	-0.74126	-0.76224	0.223776	0.454545
Paraprevo	0.380291	-0.40846	0.302824	-0.53522	0.661988	0.647903	-0.21517	-0.55635	-0.22575	-0.45776	-0.71833	-0.35212	-0.79579	0.429588	0.225358
Prevotella	-0.71329	0.132867	-0.63636	-0.08392	-0.58042	-0.78322	0.021016	0.615385	0.847637	0.769231	0.755245	0.552448	0.692308	-0.51748	-0.42657
Prevotella	0.34965	-0.58741	0.398601	-0.44056	0.601399	0.594406	-0.36778	-0.41958	-0.51489	-0.42657	-0.66434	-0.25874	-0.62937	0.230769	0.153846
Rikenellac	0.693521	0.003503	0.567426	-0.21016	0.5324	0.444834	0.463158	-0.71454	-0.17544	-0.45184	-0.46585	-0.5324	-0.59194	0.644484	0.423819
Parabacte	0.468531	0.195804	0.559441	-0.18881	0.398601	0.412587	-0.15762	-0.1958	-0.54991	-0.45455	-0.2028	-0.48252	-0.44056	0.06993	0.272727
Bilophila	0.772517	0.395077	0.72666	-0.12346	0.511484	0.419769	0.130745	-0.26809	-0.78447	-0.60673	-0.32806	-0.44446	-0.58203	0.310418	0.37744
Faecalibac	-0.6122	0.074745	-0.79728	-0.04271	-0.42355	-0.51966	-0.11766	0.701179	0.39399	0.491181	0.484062	0.647789	0.612197	-0.2883	-0.74389
Enterococ	-0.73527	0.126899	-0.7502	0.264995	-0.89949	-0.83231	-0.22994	0.824845	0.605697	0.899492	0.705411	0.843507	0.783789	-0.41429	-0.66062
Lactobacil	-0.67133	0.006993	-0.7972	0.258741	-0.48252	-0.48252	-0.36427	0.573427	0.469353	0.48951	0.272727	0.587413	0.496503	-0.02797	-0.6014
Lactococc	-0.60045	0.280728	-0.62384	0.483476	-0.81879	-0.81099	-0.04296	0.717416	0.390583	0.686224	0.733012	0.647234	0.85778	-0.3899	-0.53026
Clostridia	0.293706	-0.23776	0.223776	-0.42657	0.636364	0.685315	-0.31524	-0.40559	-0.2627	-0.4965	-0.65035	-0.36364	-0.69231	0.377622	0.223776
Clostridiur	-0.84218	-0.28853	-0.7876	0.101374	-0.68622	-0.51467	-0.06249	0.569254	0.742108	0.655032	0.436688	0.600446	0.58485	-0.53806	-0.59265
Faecalibac	-0.48677	0.653192	-0.47845	0.27043	-0.61159	-0.79049	0.297994	0.628229	0.554311	0.524218	0.81545	0.432688	0.678155	-0.52006	-0.18306
Dialister	-0.76513	0.201546	-0.74273	0.070914	-0.80245	-0.81738	-0.08786	0.757663	0.777686	0.824845	0.757663	0.694214	0.67182	-0.57851	-0.56358
Haliangiur	0.354388	0.031579	0.424564	-0.10176	0.249124	0.375441	0.442883	-0.46667	-0.2232	-0.44562	0.077193	-0.74737	-0.09123	-0.18597	0.217545
Sphingom	0.392295	0.021016	0.479861	-0.06305	0.206655	0.413311	0.477193	-0.5359	-0.10877	-0.40631	-0.03853	-0.73555	-0.23468	-0.06305	0.325745
Delftia	-0.78162	0.048617	-0.76292	0.396418	-0.91625	-0.79284	-0.01873	0.732998	0.640625	0.800315	0.70308	0.718039	0.848932	-0.50113	-0.6208
Escherichi	-0.73962	0.449571	-0.71786	0.094265	-0.70336	-0.84838	-0.09806	0.870137	0.56658	0.717863	0.913643	0.667105	0.82663	-0.73962	-0.47132
Akkerman	-0.53846	0.160839	-0.48951	-0.01399	-0.44056	-0.65035	-0.32224	0.587413	0.283713	0.615385	0.776224	0.482517	0.818182	-0.55245	-0.39161

