

Variations in diet cause alterations in microbiota and metabolites that follow changes in disease severity in a multiple sclerosis model

J.E. Libbey, J.M. Sanchez, D.J. Doty, J.T. Sim, M.F. Cusick, J.E. Cox, K.F. Fischer, J.L. Round and R.S. Fujinami

Figure S1. Flow chart detailing the treatment groups. Above the dotted line shows the animals on the different diets. Below the dotted line shows the *Lactobacillus paracasei* pre-treatment/treatment experiment.

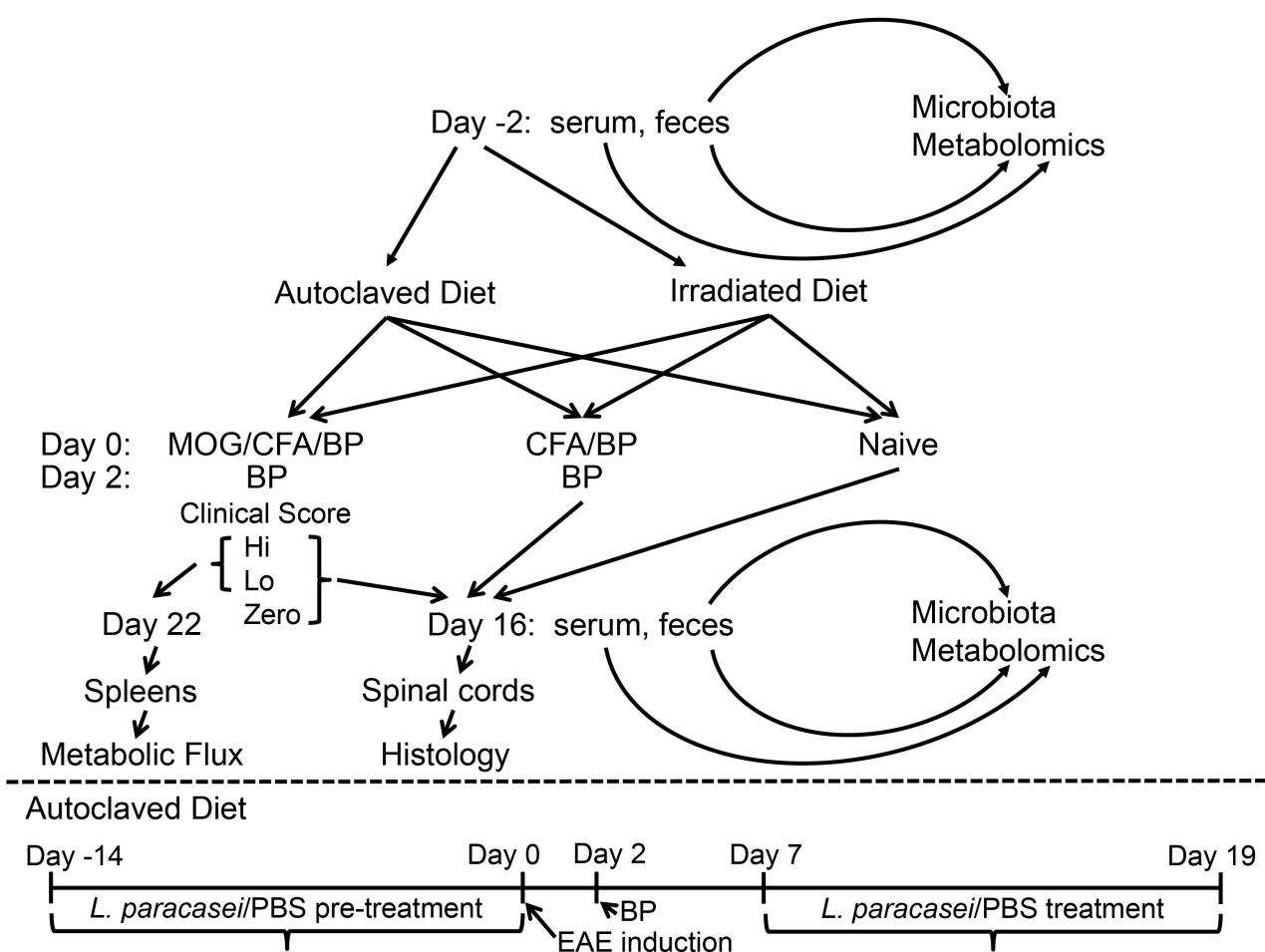
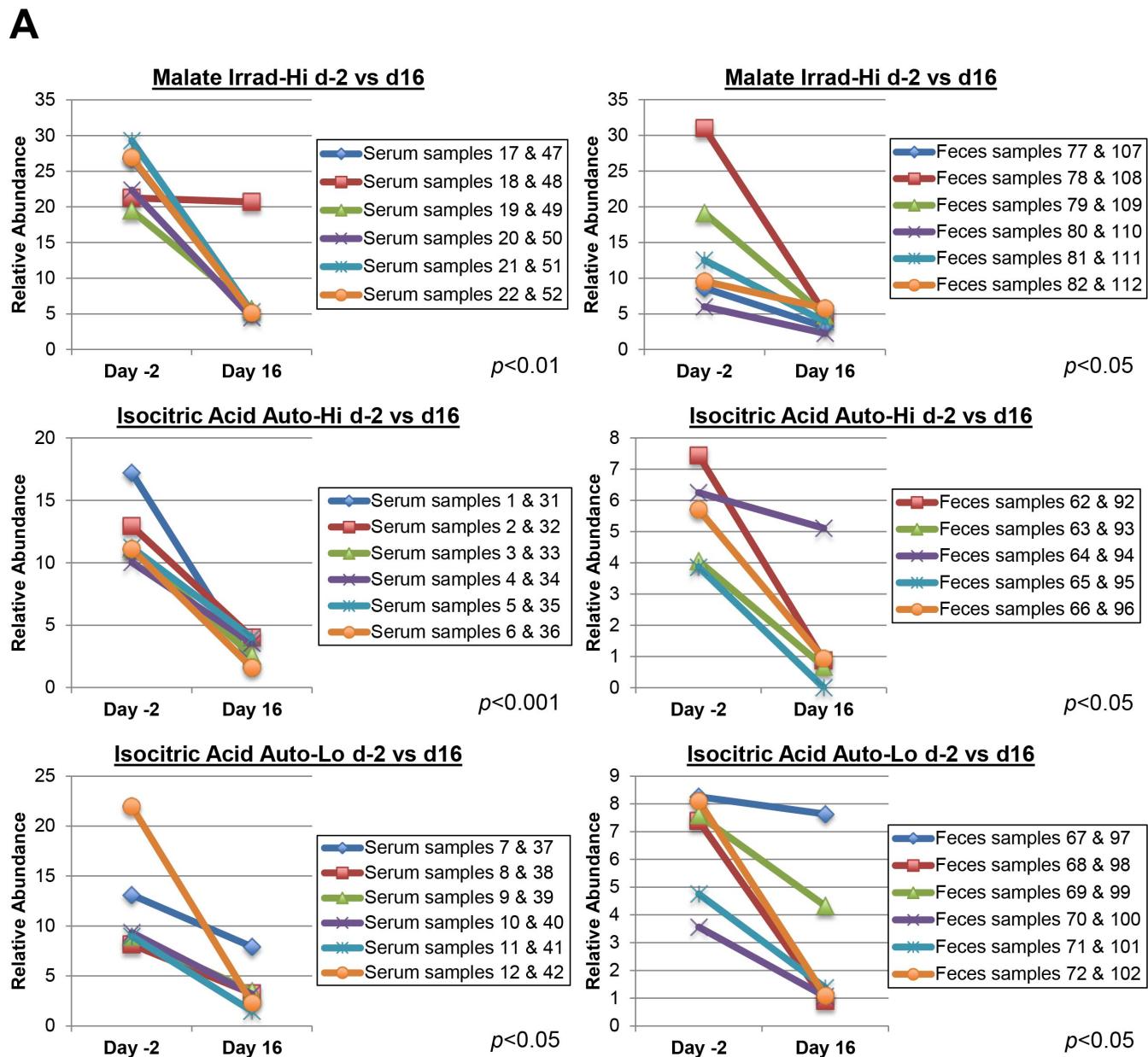


Figure S2. Correlation analyses of select metabolites. Metabolites were chosen from Figures 3 & 4 that showed the same significance for the same comparison for both serum and feces. (A) Cross-time point comparisons. Two-tailed *t* test. (B) Comparisons examining the effects of diet on disease. Spearman's rank correlation.



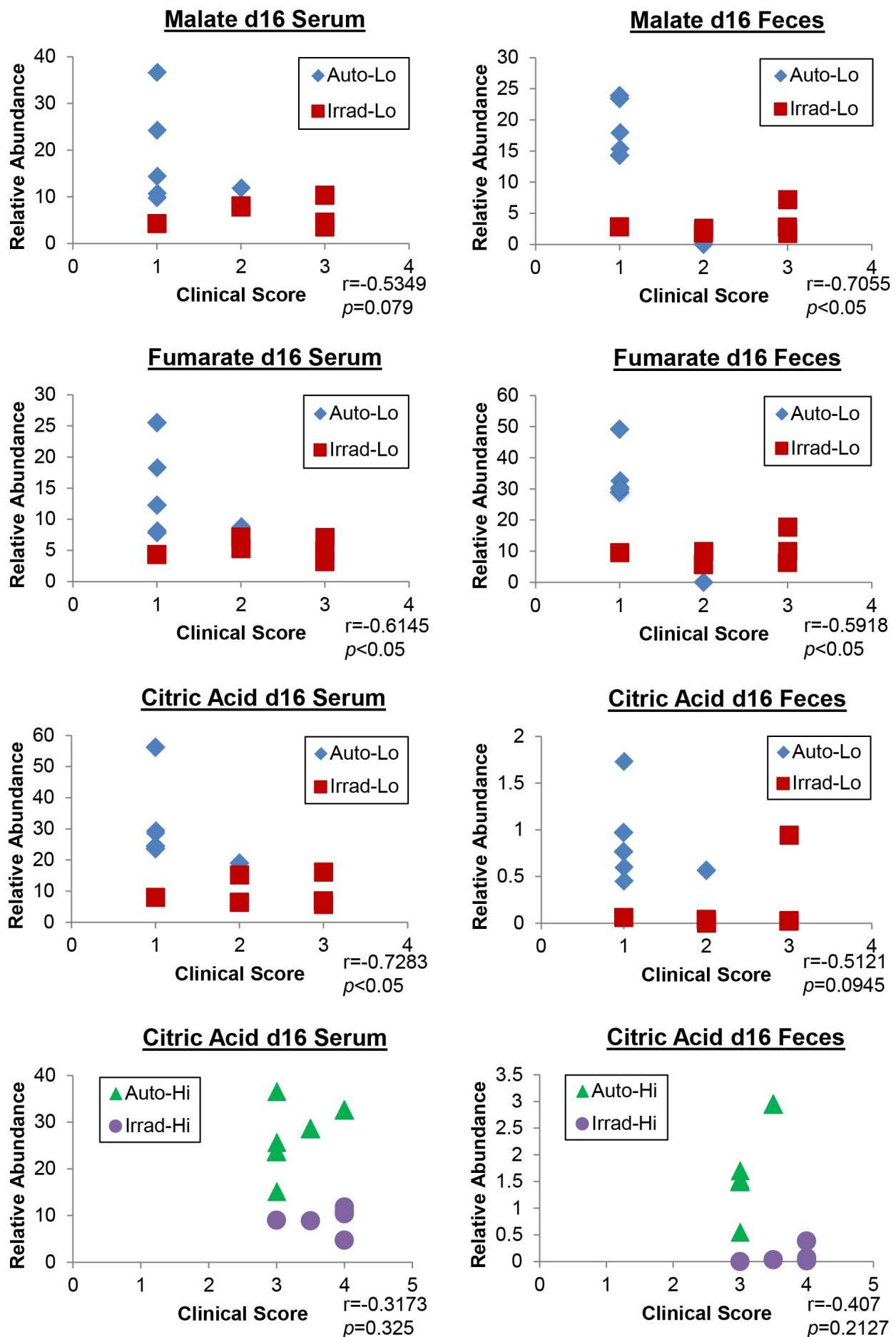
B

Table S1. Grouped x-timept comparisons day -2 vs day 16 with mice with clinical score 0.

1) Aut Zero, d-2 vs d16		2) Zero, d-2 vs d16	
Serum	Feces	Serum	Feces
1-monostearylglycerol	1-monostearylglycerol	1-monostearylglycerol	1-monostearylglycerol
1-monopalmitoylglycerol	1-monopalmitoylglycerol	1-monopalmitoylglycerol	1-monopalmitoylglycerol
4-hydroxyproline	4-hydroxyproline	4-hydroxyproline	4-hydroxyproline
homoserine	homoserine	homoserine	homoserine
isocitric acid	isocitric acid	isocitric acid	L-pipecolic acidN
succinic acid	succinic acid	succinic acid	chlorogenic acid
2-hydroxybutyric acid	2-hydroxybutyric acid	2-hydroxybutyric acid	pantothenic acid
hypoxanthine	hypoxanthine	2-ketoglutaric acid	hypoxanthine
xanthine	xanthine	uric acid	xanthine
tryptophan	tryptophan	uracil	23:0 FA
1-monooleylglycerol	lauric acid	1-monooleylglycerol	lauric acid
citric acid	un20.83_333.1338	citric acid	un20.83_333
glucose-6-phosphate	un20.83_333	glucose-6-phosphate	un18.55_204.1001
3-hydroxyanthranilic acid	un18.55_204.1001	3-hydroxyanthranilic acid	2-DOG
2-DOG	B-hydroxybutyric acid	2-DOG	un30.06_83
un30.06_83	GalNAc	un30.06_83	ribose(xyl)-5-phosphate
ribose(xyl)-5-phosphate	GlcNAc	ribose(xyl)-5-phosphate	diphosphate
diphosphate	un14.96_133.0501	diphosphate	3-(3-hydroxyphenyl)propanoic acid
3-(3-hydroxyphenyl)propanoic acid	2-hydroxyphenylpropionic acid	3-(3-hydroxyphenyl)propanoic acid	5-aminovaleric acid
1-phosphoglycerol	5-aminovaleric acid	5-aminovaleric acid	un30.81_387
urea	un30.81_387	urea	AMP
	AMP		deoxyinositol
	deoxyinositol		palmitelaidic acid
	palmitelaidic acid		B-alanine
	B-alanine		glucose
	glucose		9H-purin-6-amine
	9H-purin-6-amine		un5.79_211.0033
	un5.79_211.0033		un6.55_174.1128
	un6.55_174.1128		D-erythro-2-pentulose
	D-erythro-2-pentulose		gluconic acid
	gluconic acid		ribose
	ribose		arabinofuranose
	arabinofuranose		ornithine
	ornithine		20:0 FA
	20:0 FA		N-methylalanine
	N-methylalanine		arachidonic acid
	arachidonic acid		dihydroxyquinone
	dihydroxyquinone		un16.50_117.0729
	un16.50_117.0729		un20.99_333.1338
	un20.99_333.1338		17:0 FA
	17:0 FA		malic acid
	malic acid		galactitol
	galactitol		sorbitol
	sorbitol		22:0 FA
	22:0 FA		palmitic acid
	palmitic acid		oleic acid
	oleic acid		phosphate
	phosphate		glutamine
	glutamine		fumaric acid
	fumaric acid		orotateN
	orotateN		m-hydroxyphenylpropionic acid
	m-hydroxyphenylpropionic acid		erythrose

Table S2. Grouped x-timept comparisons day -2 vs day 16 with mice having any disease (HiLo).

1) Aut HiLo: d-2 vs d16		2) Irr HiLo: d-2 vs d16		3) HiLo: d-2 vs d16	
Serum	Feces	Serum	Feces	Serum	Feces
uracil	uracil	uracil	uracil	uracil	uracil
homoserine	homoserine	homoserine	homoserine	homoserine	homoserine
4-hydroxyproline	arachidonic acid	4-hydroxyproline	4-hydroxyproline	4-hydroxyproline	4-hydroxyproline
2-hydroxybutyric acid	2-hydroxybutyric acid	2-hydroxybutyric acid		2-hydroxybutyric acid	2-hydroxybutyric acid
glucose-6-phosphate	glucose-6-phosphate	glucose-6-phosphate		glucose-6-phosphate	glucose-6-phosphate
sedoheptulose-7-P	sedoheptulose-7-P	sedoheptulose-7-P		sedoheptulose-7-P	sedoheptulose-7-P
succinic acid	succinic acid	succinic acid		succinic acid	succinic acid
isocitric acid	isocitric acid	isocitric acid		isocitric acid	isocitric acid
2-monostearylglycerol		2-monostearylglycerol	2-monostearylglycerol	2-monostearylglycerol	
		pantothenic acid	pantothenic acid	pantothenic acid	pantothenic acid
		B-alanine	B-alanine	B-alanine	
		aconitate	aconitate	aconitate	
diphosphate	GalNAc	diphosphate	GalNAc	diphosphate	GalNAc
uric acid	GlcNAc	uric acid	GlcNAc	uric acid	GlcNAc
2-ketoglutaric acid	5-aminovaleric acid	2-ketoglutaric acid	5-aminovaleric acid	2-ketoglutaric acid	5-aminovaleric acid
citric acid	un30.81_387	citric acid	un30.81_387	citric acid	un30.81_387
3-hydroxyanthranilic acid	xanthine	3-hydroxyanthranilic acid	xanthine	3-hydroxyanthranilic acid	xanthine
N-acetylglycine	hypoxanthine	N-acetylglycine	hypoxanthine	N-acetylglycine	hypoxanthine
2-DOG	un16.50_117.0729	1-phosphoglycerol	un16.50_117.0729	1-phosphoglycerol	un16.50_117.0729
urea	un5.79_211.0033	fumaric acid	un5.79_211.0033	fumaric acid	un5.79_211.0033
linoleic acid	AMP	un30.06_83	AMP	2-DOG	AMP
	glucose	3-(3-hydroxyphenyl)propanoic	glucose	urea	glucose
	6-phosphogluconate	inositol	6-phosphogluconate		6-phosphogluconate
	phosphate	malic acid	phosphate		phosphate
	D-erythro-2-pentulose?	nicotinamide	D-erythro-2-pentulose?		D-erythro-2-pentulose?
	22:0 FA	glutamic acid	22:0 FA		22:0 FA
	1-monolinoleylglycerol		1-monolinoleylglycerol		1-monolinoleylglycerol
	ribose		ribose		ribose
	mannose		mannose		mannose
	mannitol		mannitol		mannitol
	24:0 FA		24:0 FA		24:0 FA
	24:1 FA		24:1 FA		24:1 FA
	fructose		fructose		fructose
	chlorogenic acid		chlorogenic acid		chlorogenic acid
	lysine		lysine		lysine
	leucine		leucine		leucine
	ornithine		ornithine		ornithine
	alanine		alanine		alanine
	isoleucine		isoleucine		isoleucine
	gluconic acid		gluconic acid		gluconic acid
	N-methylalanine		N-methylalanine		N-methylalanine
	un18.55_204.1001		un18.55_204.1001		un18.55_204.1001
	un14.96_133.0501		un14.96_133.0501		un14.96_133.0501
	un6.55_174.1128		L-pipecolic acidN		L-pipecolic acidN
	un20.83_333		Indolecarboxylic acid?		Indolecarboxylic acid?
	un20.83_333.1338		methionine		methionine
	9H-purin-6-amine?		glycerol		glycerol
	arabinofuranose		aspartic acid		aspartic acid
	palmitelaidic acid		N-acetylaspartate		N-acetylaspartate
	3-hydroxybutyrate		threonine		threonine
	B-hydroxybutyric acid		19:0 FA		19:0 FA
	un20.99_333.1338		thymine		thymine
	N-acetylglutamate		valine		valine
	un30.94_387.1141		sarcosine		sarcosine

ergosterol
lauric acid
pyridine-2-6-dicarboxylic acid
orotateN
1-monostearylglycerol
rhamnose
1-monopalmitoylglycerol
2,3-pyridinedicarboxylic acid
Trans-cinnamic acid
Xylose or isomer
Fructose-6-phosphate
2-phosphoglycerol
Dihydroxyquinone
un12.13_263
p-hydroxyphenylacetic acid
deoxy-inositol?
un13.58_204.1001

un13.58_204.1001
inositol
malic acid
histidine
1-monooleoylglycerol
Ribulose(xylulose)-5-P
Erythrose-4-phosphate
Mevalonic acid

un6.55_174.1128
un20.83_333
un20.83_333.1338
9H-purin-6-amine?
arabinofuranose
palmitelaidic acid
3-hydroxybutyrate
B-hydroxybutyric acid
un20.99_333.1338
N-acetylglutamate
un30.94_387.1141
ergosterol
lauric acid
pyridine-2-6-
dicarboxylic acid
orotateN
1-monostearylglycerol
rhamnose
1-monopalmitoylglycerol
2,3-pyridinedicarboxylic acid
Trans-cinnamic acid
Xylose or isomer
Fructose-6-phosphate
2-phosphoglycerol
Dihydroxyquinone
un12.13_263
p-hydroxyphenylacetic acid
deoxy-inositol?
tyrosine

Table S3. Grouped x-timept comparisons day -2 vs day 16 for different clinical score groups.

1) Hi: d-2 vs d16		2) HiLo: d-2 vs d16		3) Lo: d-2 vs d16		4) Zero: d-2 vs d16	
Serum	Feces	Serum	Feces	Serum	Feces	Serum	Feces
homoserine	homoserine	homoserine	homoserine	homoserine	homoserine	homoserine	homoserine
uracil	uracil	uracil	uracil	uracil	uracil	uracil	uracil
2-hydroxybutyric a	2-hydroxybutyric a	2-hydroxybutyric a	2-hydroxybutyric a	2-hydroxybutyric a	2-hydroxybutyric a	2-hydroxybutyric a	2-hydroxybutyric a
4-hydroxyproline	4-hydroxyproline	4-hydroxyproline	4-hydroxyproline	4-hydroxyproline	4-hydroxyproline	4-hydroxyproline	4-hydroxyproline
glucose-6-phosphate	glucose-6-phosphate	glucose-6-phosphate	glucose-6-phosphate	glucose-6-phosphate	glucose-6-phosphate	glucose-6-phosphate	glucose-6-phosphate
pantothenic acid	pantothenic acid	pantothenic acid	pantothenic acid	pantothenic acid	pantothenic acid	pantothenic acid	pantothenic acid
	1-monostearylglycerol		1-monostearylglycerol	1-monostearylglycerol	1-monostearylglycerol	1-monostearylglycerol	1-monostearylglycerol
sedoheptulose-7-P	sedoheptulose-7-P	sedoheptulose-7-P	sedoheptulose-7-P	sedoheptulose-7-P	23:0 FA	1-monopalmitoylglycerol	1-monopalmitoylglycerol
ribose(xyl)-5-P	arachidonic acid	arachidonic acid	arachidonic acid	arachidonic acid	arachidonic acid	ribose(xyl)-5-P	23:0 FA
B-alanine	B-alanine	B-alanine		B-alanine			B-alanine
succinic acid		succinic acid	succinic acid	succinic acid		succinic acid	
3-(3-hydroxyphenyl) propanoic acid	2-hydroxyphenylpropionic acid	3-(3-hydroxyphenyl) propanoic acid	deoxy-inositol?	3-(3-hydroxyphenyl) propanoic acid	3-(3-hydroxyphenyl) propanoic acid	3-(3-hydroxyphenyl) propanoic acid	2-hydroxyphenylpropionic acid
3-hydroxyanthranilic a	GalNAc	3-hydroxyanthranilic a	GalNAc	3-hydroxyanthranilic a	GalNAc	3-hydroxyanthranilic a	GalNAc
2-DOG	GlcNAc	2-DOG	GlcNAc	2-DOG	GlcNAc	2-DOG	GlcNAc
un30.06_83	5-aminovaleric acid	un30.06_83	5-aminovaleric acid	un30.06_83	5-aminovaleric acid	un30.06_83	5-aminovaleric acid
diphosphate	xanthine	diphosphate	xanthine	diphosphate	xanthine	diphosphate	xanthine
2-ketoglutaric acid	un20.83_333.1338	2-ketoglutaric acid	un20.83_333.1338	2-ketoglutaric acid	un20.83_333.1338	2-ketoglutaric acid	un20.83_333.1338
uric acid	un20.83_333	uric acid	un20.83_333	uric acid	un20.83_333	uric acid	un20.83_333
isocitric acid	un14.96_133.0501	isocitric acid	un14.96_133.0501	isocitric acid	un14.96_133.0501	isocitric acid	un14.96_133.0501
citric acid	un30.81_387	citric acid	un30.81_387	citric acid	un30.81_387	citric acid	un30.81_387
N-acetylglycine	AMP	N-acetylglycine	AMP	N-acetylglycine	AMP	N-acetylglycine	AMP
1-phosphoglycerol	L-pipecolic acidN	1-phosphoglycerol	L-pipecolic acidN	1-phosphoglycerol	L-pipecolic acidN	1-phosphoglycerol	L-pipecolic acidN
fumaric acid	palmitelaidic acid	fumaric acid	palmitelaidic acid	fumaric acid	palmitelaidic acid	fumaric acid	palmitelaidic acid
urea	glucose	urea	glucose	urea	glucose	urea	glucose
2-monostearylglycerol	9H-purin-6-amine	2-monostearylglycerol	9H-purin-6-amine	2-monostearylglycerol	9H-purin-6-amine	2-monostearylglycerol	9H-purin-6-amine
linoleic acid	un18.55_204.1001	aconitate	un18.55_204.1001	homocysteine	un18.55_204.1001	homocysteine	un18.55_204.1001
	N-methylalanine		N-methylalanine		N-methylalanine		N-methylalanine
	gluconic acid		gluconic acid		gluconic acid		gluconic acid
	ribose		ribose		ribose		ribose
	hypoxanthine		hypoxanthine		hypoxanthine		hypoxanthine
	D-erythro-2-pentulose		D-erythro-2-pentulose		D-erythro-2-pentulose		D-erythro-2-pentulose
	un5.79_211.0033		un5.79_211.0033		un5.79_211.0033		un5.79_211.0033
	arabinofuranose		arabinofuranose		arabinofuranose		arabinofuranose
	ornithine		ornithine		ornithine		ornithine
	un6.55_174.1128		un6.55_174.1128		un6.55_174.1128		un6.55_174.1128
	lauric acid		lauric acid		lauric acid		lauric acid
	1-monooleoylglycerol		chlorogenic acid		chlorogenic acid		chlorogenic acid
	B-hydroxybutyric acid		B-hydroxybutyric acid		B-hydroxybutyric acid		B-hydroxybutyric acid
	6-phosphogluconate		6-phosphogluconate		6-phosphogluconate		6-phosphogluconate
	pyridine-2-6-dicarboxylic a		pyridine-2-6-dicarboxylic a		pyridine-2-6-dicarboxylic a		pyridine-2-6-dicarboxylic a
	24:0 FA		24:0 FA		24:0 FA		24:0 FA
	24:1 FA		24:1 FA		24:1 FA		24:1 FA
	fructose		fructose		fructose		fructose
	mannose		mannose		mannose		mannose
	mannitol		mannitol		mannitol		mannitol
	22:0 FA		22:0 FA		22:0 FA		22:0 FA
	alanine		alanine		alanine		alanine
	orotateN		orotateN		orotateN		orotateN
	N-acetylglutamate		N-acetylglutamate		N-acetylglutamate		N-acetylglutamate
	rhamnose		rhamnose		rhamnose		rhamnose
	ergosterol		ergosterol		ergosterol		ergosterol
	sarcosine		sarcosine		sarcosine		sarcosine
	1-monolinoleoylglycerol		1-monolinoleoylglycerol		1-monolinoleoylglycerol		1-monolinoleoylglycerol
	un16.50_117.0729		un16.50_117.0729		un16.50_117.0729		un16.50_117.0729
	3-hydroxybutyrate		3-hydroxybutyrate		3-hydroxybutyrate		3-hydroxybutyrate
	N-acetylaspartate		N-acetylaspartate		N-acetylaspartate		N-acetylaspartate
	phosphate		phosphate		phosphate		phosphate

un20.99_333.1338
un30.94_387.1141
thymine
Dihydroxyquinone
trans-cinnamic acid
indolecarboxylic acid?
un12.13_263
2,3-pyridinedicarboxylic a
fructose-6-phosphate
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2,3-pyridinedicarboxylic a
fructose-6-phosphate

aminomalonic acid
ribulose(xylulose)-5-P
erythrose-4-phosphate
sorbitol
3-phosphoglycerate

leucine
isoleucine
methionine
glycerol
lysine
aspartic acid
threonine
19:0 FA
valine
Xylose or isomer
2-phosphoglycerol
p-hydroxyphenylacetic a
tyrosine

leucine
isoleucine
methionine
glycerol
lysine
aspartic acid
threonine
19:0 FA
valine
Xylose or isomer
2-phosphoglycerol
p-hydroxyphenylacetic a
tyrosine