

### Supplemental Digital Content

**Supplemental Figure 1. A shift in plasma metabolomic profiles in HF and after resynchronization therapy (CRT). A, A three way comparison of plasma metabolomic profiles of control, HF pre-CRT and HF post CRT groups by partial least squares discriminant analysis (PLS-DA). B, An integral panel of the most important metabolites in group distinction by variable importance in the projection (VIP).**

**Supplemental Figure 2.** Transitions in metabolomic matrix in control patients, and cardiac resynchronization therapy (CRT) patients. The red color in this heat map indicates an increased metabolite level, and green indicates a decreased metabolite level. The CRT group has a distinct cluster (#1) with decreased metabolite levels and another cluster (#2 and below) with increased metabolite levels. Glutamic acid and glutamic acid (dehydrated) are shown as two separate entries.

Supplemental Table 1. Plasma Metabolite Levels Before and After CRT\*

Metabolite	Control			HF pre-CRT (N=24)			HF post-CRT (N=18)			HF pre-CRT – Control P Value	HF post-CRT – Control P Value	HF pre-CRT- post-CRT P Value
2-HB	1.44	±	0.35	1.14	±	0.11	0.60	±	0.12	.29	.010	.002
2-hydroxypyridine	0.82	±	0.10	0.89	±	0.07	1.23	±	0.11	.63	.020	.009
3-AIB	0.78	±	0.13	0.89	±	0.08	1.25	±	0.12	.51	.023	.01
3-HB	0.85	±	0.09	0.93	±	0.06	1.17	±	0.10	.51	.052	.04
Acetate	0.71	±	0.18	0.78	±	0.12	1.43	±	0.19	.76	.023	.006
Acetoacetate	0.80	±	0.12	0.90	±	0.10	1.23	±	0.13	.55	.040	.04
Acetone	0.80	±	0.23	0.96	±	0.09	1.15	±	0.12	.42	.14	.22
Acetylsalicylic acid	0.13	±	0.07	1.24	±	0.54	1.15	±	0.45	.19	.11	.90
Alanine	0.92	±	0.16	0.78	±	0.10	1.32	±	0.16	.49	.12	.006
Allo-inositol	0.51	±	0.09	0.97	±	0.19	1.29	±	0.21	.13	.015	.26
Arabitol	0.53	±	0.11	0.98	±	0.17	1.28	±	0.25	.11	.047	.31
Benzoate	1.49	±	0.47	1.18	±	0.33	0.51	±	0.07	.61	.010	.09
Beta-alanine	0.95	±	0.13	0.92	±	0.07	1.13	±	0.10	.83	.28	.08
Cholesterol	1.15	±	0.14	0.90	±	0.05	1.05	±	0.07	.04	.46	.09
Citrate	1.08	±	0.26	0.85	±	0.07	1.15	±	0.21	.26	.002	.15
Creatine	0.75	±	0.07	0.91	±	0.06	1.24	±	0.09	.12	.001	.003
Creatinine	0.58	±	0.10	1.13	±	0.17	1.06	±	0.15	.05	.002	.79
Cysteine	1.26	±	0.19	0.83	±	0.08	1.07	±	0.07	.02	.27	.04
Fumarate	0.60	±	0.12	1.01	±	0.18	1.20	±	0.17	.16	.021	.45
Galactinol	0.51	±	0.12	0.46	±	0.08	1.93	±	0.47	.75	.038	.001
Galacturonate	0.52	±	0.17	1.85	±	0.48	0.18	±	0.00	.09	.007	.004
Gluconate	0.76	±	0.28	1.73	±	0.43	0.21	±	0.00	.17	.009	.003
Glucose	0.91	±	0.08	0.98	±	0.02	1.07	±	0.05	.24	.099	.12
Glutamate	0.43	±	0.08	1.54	±	0.14	0.61	±	0.09	<.001	.21	<.001
Glutamine	0.83	±	0.14	0.81	±	0.11	1.33	±	0.15	.900	.042	.007
Glycerate	0.62	±	0.10	1.16	±	0.19	1.00	±	0.12	.09	.040	.52
Glycerol	1.92	±	0.41	0.91	±	0.14	0.63	±	0.09	.005	.000	.13
Glycerol-1- phosphate	1.01	±	0.30	0.79	±	0.07	1.27	±	0.12	.32	.34	.001
Glycine	1.29	±	0.22	0.82	±	0.05	1.07	±	0.12	.007	.36	.04
Glycolate	0.89	±	0.07	0.93	±	0.07	1.14	±	0.08	.75	.052	.05
Heptadecanoate	1.80	±	0.80	1.18	±	0.19	0.35	±	0.00	.30	.018	<.001
Iminodiacetate	0.94	±	0.05	1.00	±	0.05	1.03	±	0.05	.47	.28	.69
Isoleucine	0.87	±	0.09	0.90	±	0.05	1.19	±	0.07	.77	.015	.001
IBTP	1.10	±	0.46	0.89	±	0.07	1.08	±	0.21	.50	.96	.37
Lactate	1.03	±	0.25	0.92	±	0.08	1.09	±	0.10	.60	.78	.19
Leucine	0.87	±	0.10	0.90	±	0.05	1.19	±	0.08	.76	.026	.004
Linoleate	1.85	±	0.51	0.94	±	0.08	0.63	±	0.08	.01	.003	.009
Lysine	0.93	±	0.07	0.97	±	0.12	1.08	±	0.16	.86	.52	.55
Malate	0.85	±	0.34	1.59	±	0.35	0.34	±	0.00	.22	.047	.003

<b>Malonate</b>	0.65	±	0.05	0.94	±	0.13	1.26	±	0.25	.16	.09	.24
<b>Mannitol</b>	1.13	±	0.56	0.87	±	0.27	1.10	±	0.36	.64	.97	.60
<b>Melezitose</b>	1.47	±	0.61	1.52	±	0.50	0.10	±	0.00	.96	.004	.02
<b>Methylamine</b>	0.61	±	0.31	0.72	±	0.21	1.57	±	0.30	.78	.049	.02
<b>Methylguanidine</b>	1.22	±	0.36	0.77	±	0.23	1.17	±	0.21	.31	.90	.22
<b>Myristate</b>	1.66	±	0.33	1.14	±	0.09	0.48	±	0.05	.05	<.001	<.001
<b>Nicotinate</b>	1.41	±	0.17	1.14	±	0.12	0.60	±	0.17	.22	.005	.01
<b>N-methylhistidine</b>	0.47	±	0.04	0.59	±	0.04	1.80	±	0.38	.07	.018	.001
<b>Norleucine</b>	0.90	±	0.10	0.94	±	0.09	1.12	±	0.13	.78	.26	.26
<b>Norvaline</b>	1.06	±	0.12	1.01	±	0.07	0.96	±	0.12	.70	.56	.69
<b>Oleate</b>	1.50	±	0.30	1.18	±	0.09	0.51	±	0.08	.20	<.001	<.001
<b>Ornithine</b>	0.95	±	0.12	0.95	±	0.07	1.09	±	0.09	.99	.34	.12
<b>Oxalate</b>	0.60	±	0.11	0.78	±	0.09	1.49	±	0.27	.26	.029	.01
<b>Palmitate</b>	1.33	±	0.19	1.08	±	0.06	0.73	±	0.05	.10	<.001	<.001
<b>Palmitoleate</b>	1.81	±	0.51	1.18	±	0.17	0.35	±	0.09	.13	<.001	<.001
<b>Phenylalanine</b>	0.59	±	0.09	0.54	±	0.05	1.79	±	0.57	.62	.14	.02
<b>Phosphate</b>	0.98	±	0.11	0.95	±	0.03	1.07	±	0.04	.71	.34	.02
<b>Proline</b>	0.76	±	0.11	0.89	±	0.08	1.26	±	0.09	.33	.002	.004
<b>Pyrophosphate</b>	0.31	±	0.09	0.72	±	0.26	1.72	±	1.02	.33	.33	.30
<b>Pyruvate</b>	1.15	±	0.34	0.98	±	0.11	0.94	±	0.16	.56	.54	.82
<b>Serine</b>	1.42	±	0.15	0.93	±	0.07	0.87	±	0.08	.003	.001	.55
<b>Serotonin</b>	1.62	±	0.18	1.06	±	0.12	0.59	±	0.15	.02	<.001	.02
<b>Stearate</b>	1.15	±	0.12	1.08	±	0.04	0.82	±	0.04	.49	.005	<.001
<b>Succinate</b>	0.90	±	0.33	0.73	±	0.14	1.40	±	0.21	.58	<.001	.009
<b>Sucrose</b>	0.06	±	0.03	0.58	±	0.39	2.03	±	1.08	.40	.20	.12
<b>Tagatose</b>	0.46	±	0.14	1.91	±	0.67	0.14	±	0.00	.18	.003	.02
<b>Threitol</b>	0.48	±	0.08	1.08	±	0.14	1.17	±	0.20	.01	.025	.71
<b>Threonine</b>	1.40	±	0.16	0.85	±	0.08	0.98	±	0.08	.002	.017	.24
<b>Tryptophan</b>	0.86	±	0.23	1.63	±	0.29	0.27	±	0.00	.11	.001	<.001
<b>Tyrosine</b>	0.49	±	0.04	0.50	±	0.03	1.90	±	0.41	.85	.021	<.001
<b>Urea</b>	0.72	±	0.08	1.03	±	0.09	1.11	±	0.12	.05	.038	.58
<b>Urate</b>	0.47	±	0.07	1.14	±	0.13	1.11	±	0.19	.003	.023	.88
<b>Valine</b>	0.93	±	0.10	0.94	±	0.05	1.10	±	0.07	.92	.031	.06

Abbreviations: CRT, cardiac resynchronization therapy; IBTP, isopropyl  $\beta$ -1-thiogalactopyranoside; 2-HB, 2-hydroxybutyrate; 3-AIB, 3-aminoisobutyrate; 3-HB, 3-hydroxybutyrate.

\* Values are mean $\pm$ standard error of the mean unless indicated otherwise.

**Supplemental Table 2.** Plasma Metabolite Levels Before and After CRT in Responders and Non-responders\*

Metabolite	Responders							Non-responders						
	pre-CRT(N=9)			post-CRT(N=7)			P Value	pre-CRT(N=13)			post-CRT(N=10)			P Value
2-HB	1.24	±	0.60	0.79	±	0.73	0.15	1.05	±	0.44	0.43	±	0.13	<.001
2-hydroxypyridine	0.75	±	0.38	1.25	±	0.51	0.02	1.01	±	0.30	1.22	±	0.44	0.19
3-AIB	0.95	±	0.40	1.24	±	0.41	0.13	0.83	±	0.41	1.27	±	0.65	0.06
3-HB	0.98	±	0.28	1.14	±	0.35	0.26	0.88	±	0.33	1.20	±	0.53	0.10
Acetate	0.78	±	0.53	1.37	±	0.72	0.05	0.78	±	0.68	1.48	±	0.96	0.05
Acetoacetate	0.94	±	0.39	1.19	±	0.51	0.23	0.86	±	0.56	1.27	±	0.64	0.12
Acetone	1.03	±	0.40	1.22	±	0.53	0.36	0.91	±	0.50	1.08	±	0.53	0.43
Acetylsalicylic acid	1.23	±	3.03	0.24	±	0.46	0.35	1.26	±	2.37	1.97	±	2.41	0.49
Alanine	0.80	±	0.45	1.33	±	0.55	0.03	0.77	±	0.58	1.30	±	0.82	0.08
Allo-inositol	1.04	±	1.13	1.53	±	1.11	0.34	0.92	±	0.74	1.07	±	0.68	0.61
Arabitol	1.02	±	0.82	1.51	±	1.46	0.35	0.94	±	0.87	1.07	±	0.64	0.70
Benzoic acid	1.07	±	0.72	0.55	±	0.35	0.05	1.27	±	2.16	0.48	±	0.29	0.26
Beta- alanine	0.86	±	0.38	1.24	±	0.58	0.10	0.96	±	0.34	1.04	±	0.27	0.55
Cholesterol	0.95	±	0.18	1.16	±	0.33	0.15	0.85	±	0.29	0.95	±	0.27	0.43
Citrate	0.90	±	0.49	1.35	±	0.58	0.08	0.76	±	0.52	1.52	±	0.82	0.01
Creatine	0.92	±	0.22	1.16	±	0.29	0.05	0.91	±	0.33	1.31	±	0.47	0.03
Creatinine	0.89	±	0.25	1.20	±	0.42	0.05	0.89	±	0.46	1.39	±	0.62	0.03
Cysteine	0.84	±	0.40	1.11	±	0.26	0.09	0.83	±	0.41	1.04	±	0.38	0.23
Fumaric acid	1.14	±	1.11	1.27	±	0.93	0.78	0.90	±	0.65	1.14	±	0.52	0.36
Galactinol	0.52	±	0.48	1.79	±	2.36	0.10	0.42	±	0.27	2.06	±	1.81	<.001
Galacturonic acid	2.21	±	2.41	0.18	±	0.00	0.02	1.54	±	2.37	0.18	±	0.00	0.09
Gluconic acid	1.87	±	2.14	0.21	±	0.00	0.03	1.61	±	2.14	0.21	±	0.00	0.05
Glucose	0.98	±	0.09	1.08	±	0.16	0.07	0.99	±	0.12	1.05	±	0.28	0.48
Glutamic acid	1.58	±	0.68	0.70	±	0.39	<.001	1.51	±	0.68	0.54	±	0.43	<.001
Glutamine	0.81	±	0.45	1.25	±	0.58	0.07	0.81	±	0.64	1.40	±	0.74	0.06
Glyceric acid	1.06	±	0.29	0.92	±	0.40	0.38	1.24	±	1.25	1.08	±	0.59	0.71
Glycerol	0.85	±	0.33	0.69	±	0.48	0.39	0.95	±	0.89	0.58	±	0.35	0.23
Glycerol -1-P	0.88	±	0.29	1.42	±	0.45	<.001	0.71	±	0.35	1.13	±	0.53	0.03
Glycine	0.75	±	0.24	0.92	±	0.36	0.22	0.88	±	0.25	1.21	±	0.62	0.10
Glycolic acid	0.88	±	0.31	1.08	±	0.35	0.18	0.98	±	0.35	1.20	±	0.36	0.15
Heptadecanoic acid	1.30	±	1.03	0.35	±	0.00	0.01	1.07	±	0.88	0.35	±	0.00	0.02
Iminodiacetic acid	0.96	±	0.14	1.02	±	0.21	0.47	1.03	±	0.28	1.03	±	0.23	0.99
Isoleucine	0.94	±	0.17	1.15	±	0.27	0.05	0.87	±	0.28	1.23	±	0.37	0.01
Isopropyl beta--thiogalactopyranoside	0.87	±	0.32	1.07	±	0.68	0.39	0.92	±	0.34	1.09	±	1.16	0.61
Lactate	0.98	±	0.42	1.21	±	0.44	0.24	0.87	±	0.42	0.98	±	0.43	0.54
Leucine	0.95	±	0.19	1.15	±	0.30	0.08	0.87	±	0.30	1.22	±	0.42	0.03
Linoleic acid	1.06	±	0.49	0.69	±	0.47	0.11	0.84	±	0.26	0.56	±	0.22	0.01
Lysine	0.97	±	0.44	1.15	±	0.65	0.47	0.96	±	0.68	1.02	±	0.76	0.85

Malic acid	1.38	±	1.60	0.34	±	0.00	0.07	1.76	±	1.89	0.34	±	0.00	0.03
Malonate	1.07	±	0.89	1.22	±	1.15	0.75	0.83	±	0.29	1.29	±	1.06	0.15
Mannitol	1.01	±	1.52	0.70	±	0.98	0.61	0.75	±	1.17	1.46	±	1.94	0.29
Melezitose	1.97	±	2.89	0.10	±	0.00	0.07	1.14	±	2.06	0.10	±	0.00	0.13
Methylamine	0.79	±	0.99	1.43	±	1.13	0.19	0.65	±	1.13	1.68	±	1.47	0.07
Methylguanidine	0.75	±	0.81	1.14	±	0.85	0.30	0.80	±	1.37	1.19	±	1.01	0.45
Myristic acid	1.05	±	0.42	0.49	±	0.21	<.001	1.21	±	0.48	0.48	±	0.26	<.001
Nicotinic acid	1.15	±	0.50	0.46	±	0.53	0.01	1.14	±	0.66	0.72	±	0.90	0.21
N-methylhistidine	0.54	±	0.14	1.37	±	1.18	0.03	0.63	±	0.22	2.18	±	1.98	0.01
Norleucine	1.07	±	0.54	1.32	±	0.76	0.41	0.84	±	0.35	0.95	±	0.28	0.41
Norvaline	1.10	±	0.36	1.16	±	0.59	0.81	0.93	±	0.36	0.78	±	0.35	0.32
Oleic acid	1.25	±	0.45	0.56	±	0.36	0.00	1.12	±	0.48	0.46	±	0.31	0.00
Ornithine	0.93	±	0.23	1.19	±	0.28	0.03	0.96	±	0.39	1.01	±	0.45	0.80
Oxalic acid	0.69	±	0.50	1.44	±	0.91	0.03	0.86	±	0.37	1.53	±	1.43	0.12
Palmitic acid	1.13	±	0.29	0.76	±	0.26	0.01	1.03	±	0.27	0.70	±	0.18	<.001
Palmitoleic acid	1.34	±	0.69	0.38	±	0.41	<.001	1.04	±	0.93	0.32	±	0.39	0.03
Phenylalanine	0.62	±	0.19	1.30	±	1.18	0.08	0.47	±	0.31	2.24	±	3.25	0.06
Phosphoric acid	0.92	±	0.20	1.08	±	0.16	0.05	0.98	±	0.13	1.06	±	0.20	0.22
Porphine	0.72	±	0.26	0.91	±	0.37	0.18	1.04	±	0.36	1.61	±	0.58	0.01
Proline	0.90	±	0.38	1.19	±	0.33	0.09	0.89	±	0.41	1.32	±	0.45	0.02
Pyrophosphate	1.07	±	1.82	1.47	±	4.13	0.77	0.42	±	0.45	1.94	±	4.91	0.28
Pyruvic acid	1.12	±	0.64	1.21	±	0.76	0.79	0.87	±	0.44	0.70	±	0.56	0.44
Serine	1.03	±	0.47	0.93	±	0.34	0.58	0.85	±	0.24	0.81	±	0.35	0.78
Serotonin	1.17	±	0.56	0.66	±	0.73	0.09	0.97	±	0.63	0.54	±	0.60	0.11
Stearic acid	1.08	±	0.20	0.85	±	0.21	0.02	1.08	±	0.17	0.79	±	0.17	0.00
Succinate	0.68	±	0.60	1.30	±	0.78	0.05	0.77	±	0.74	1.49	±	1.07	0.07
Sucrose	0.94	±	2.78	3.75	±	6.57	0.21	0.27	±	0.54	0.49	±	0.89	0.49
Tagatose	1.26	±	1.38	0.14	±	0.00	0.03	2.45	±	4.26	0.14	±	0.00	0.10
Threitol	1.08	±	0.59	1.38	±	1.16	0.46	1.08	±	0.80	0.98	±	0.56	0.74
Threonine	0.84	±	0.38	1.02	±	0.31	0.27	0.86	±	0.38	0.94	±	0.40	0.60
Tryptophan	1.59	±	1.36	0.27	±	0.00	0.01	1.67	±	1.50	0.27	±	0.00	0.01
Tyrosine	0.50	±	0.12	1.42	±	1.32	0.03	0.50	±	0.16	2.34	±	2.13	0.01
Urea	1.07	±	0.42	1.14	±	0.59	0.76	0.99	±	0.48	1.08	±	0.50	0.65
Uric acid	1.21	±	0.51	1.37	±	1.04	0.68	1.07	±	0.73	0.87	±	0.50	0.46
Valine	0.94	±	0.24	1.18	±	0.32	0.07	0.85	±	0.35	1.24	±	0.46	0.03

Abbreviations: CRT, cardiac resynchronization therapy; IBTP, isopropyl  $\beta$ -1-thiogalactopyranoside; 2-HB, 2-hydroxybutyrate; 3-AIB, 3-aminoisobutyrate; 3-HB, 3-hydroxybutyrate.

\* Values are mean $\pm$ standard error of the mean unless indicated otherwise.

**Supplemental Table 3.** Changes (delta) in Plasma Metabolite Levels Before and After CRT in Responders and Nonresponders\*

Metabolite	Responders post-CRT- pre-CRT (N=7)			Nonresponders post-CRT- pre-CRT (N=10)			P Value
		±			±		
2-HB	-0.62	±	0.93	-0.62	±	0.39	0.98
2-hydroxypyridine	0.45	±	0.61	0.20	±	0.43	0.33
3-AIB	0.27	±	0.64	0.37	±	0.60	0.74
3-HB	0.13	±	0.49	0.24	±	0.42	0.61
Acetate	0.44	±	0.95	0.54	±	1.11	0.84
Acetoacetate	0.13	±	0.58	0.27	±	0.78	0.68
Acetone	0.14	±	0.71	0.16	±	0.43	0.94
Acetylsalicylic acid	-1.18	±	3.70	0.45	±	3.02	0.32
Alanine	0.44	±	0.78	0.46	±	0.96	0.96
Allo-inositol	0.44	±	0.73	0.10	±	0.51	0.26
Arabitol	0.46	±	0.98	0.05	±	0.57	0.28
Benzoic acid	-0.62	±	0.69	-1.00	±	2.59	0.70
Beta- alanine	0.31	±	0.66	0.08	±	0.56	0.42
Cholesterol	0.15	±	0.18	0.09	±	0.14	0.43
Citrate	0.43	±	0.67	0.61	±	0.78	0.62
Creatine	0.18	±	0.30	0.29	±	0.50	0.57
Creatinine	0.25	±	0.43	0.36	±	0.70	0.69
Cysteine	0.29	±	0.39	0.15	±	0.56	0.57
Fumaric acid	-0.04	±	1.17	0.14	±	0.30	0.64
Galactinol	1.49	±	2.52	1.63	±	1.84	0.90
Galacturonic acid	-2.40	±	2.73	-1.45	±	2.59	0.46
Gluconic acid	-2.02	±	2.38	-1.41	±	2.40	0.60
Glucose	0.08	±	0.09	0.05	±	0.23	0.78
Glutamic acid	-0.80	±	0.73	-1.06	±	0.66	0.44
Glutamine	0.30	±	0.70	0.45	±	0.94	0.71
Glyceric acid	-0.10	±	0.33	-0.35	±	1.22	0.58
Glycerol	-0.22	±	0.45	-0.50	±	0.75	0.37
Glycerol -1-P	0.38	±	0.43	0.47	±	0.41	0.67
Glycine	0.21	±	0.17	0.32	±	0.48	0.56
Glycolic acid	0.36	±	0.42	0.26	±	0.48	0.65
Heptadecanoic acid	-1.05	±	1.04	-0.76	±	0.96	0.55
Iminodiacetic acid	0.04	±	0.22	-0.06	±	0.30	0.41
Isoleucine	0.18	±	0.37	0.31	±	0.38	0.47
Isopropyl beta--thiogalactopyranoside	-0.03	±	0.25	0.25	±	1.08	0.49
Lactate	0.18	±	0.56	0.14	±	0.40	0.88
Leucine	0.18	±	0.41	0.30	±	0.41	0.55
Linoleic acid	-0.46	±	0.29	-0.33	±	0.22	0.29

Lysine	0.24	±	0.63	0.10	±	1.08	0.74
Malic acid	-1.18	±	1.79	-1.44	±	2.08	0.78
Malonate	0.03	±	1.30	0.46	±	1.04	0.44
Mannitol	-0.48	±	2.13	0.65	±	2.23	0.29
Melezitose	-2.08	±	3.29	-1.07	±	2.25	0.45
Methylamine	0.43	±	1.82	0.84	±	1.72	0.63
Methylguanidine	0.19	±	1.37	0.16	±	1.94	0.97
Myristic acid	-0.67	±	0.52	-0.87	±	0.55	0.46
Nicotinic acid	-0.70	±	0.58	-0.40	±	1.46	0.59
N-methylhistidine	0.88	±	1.20	1.54	±	1.95	0.42
Norleucine	0.47	±	0.74	0.10	±	0.47	0.21
Norvaline	0.14	±	0.65	-0.13	±	0.24	0.25
Oleic acid	-0.84	±	0.44	-0.77	±	0.42	0.72
Ornithine	0.37	±	0.32	-0.06	±	0.46	0.04**
Oxalic acid	0.76	±	0.86	0.70	±	1.37	0.91
Palmitic acid	-0.48	±	0.33	-0.39	±	0.24	0.47
Palmitoleic acid	-1.20	±	0.72	-0.91	±	0.86	0.46
Phenylalanine	0.62	±	1.27	1.69	±	3.15	0.38
Phosphoric acid	0.22	±	0.10	0.09	±	0.22	0.13
Porphine	0.10	±	0.46	0.63	±	0.47	0.03**
Proline	0.47	±	0.49	0.35	±	0.48	0.61
Pyrophosphate	0.29	±	5.23	1.47	±	5.10	0.64
Pyruvic acid	-0.02	±	0.53	-0.09	±	0.59	0.79
Serine	0.07	±	0.32	-0.08	±	0.29	0.30
Serotonin	-0.30	±	0.88	-0.48	±	1.15	0.72
Stearic acid	-0.28	±	0.35	-0.31	±	0.17	0.76
Succinate	0.42	±	0.95	0.56	±	1.24	0.80
Sucrose	2.93	±	6.55	0.16	±	1.17	0.21
Tagatose	-0.97	±	1.07	-2.25	±	4.80	0.47
Threitol	0.29	±	0.80	-0.20	±	0.54	0.14
Threonine	0.32	±	0.29	0.03	±	0.34	0.08
Tryptophan	-1.01	±	1.01	-1.45	±	1.66	0.52
Tyrosine	1.02	±	1.30	1.85	±	2.16	0.36
Urea	0.06	±	0.36	0.04	±	0.26	0.90
Uric acid	0.23	±	0.88	-0.32	±	0.43	0.10
Valine	0.22	±	0.47	0.34	±	0.49	0.60

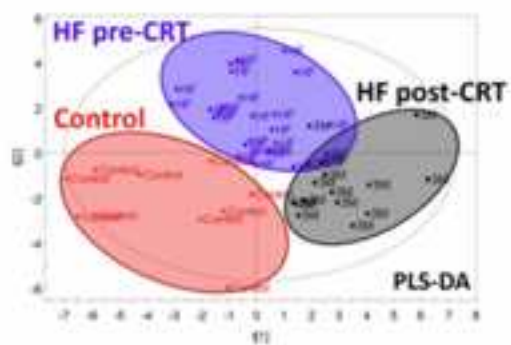
Abbreviations: CRT, cardiac resynchronization therapy; IBTP, isopropyl  $\beta$ -1-thiogalactopyranoside; 2-HB, 2-hydroxybutyrate; 3-AIB, 3-aminoisobutyrate; 3-HB, 3-hydroxybutyrate.

\* Values are mean $\pm$ standard error of the mean unless indicated otherwise.

\*\*  $p$  Value <.05.

Figure 1S

**A** Discrimination of metabolomic profiles



**B** Most important metabolites in group distinction

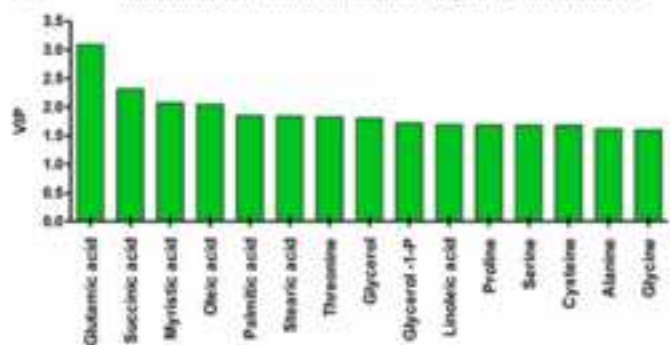




Figure 2S

