

Fig. 1

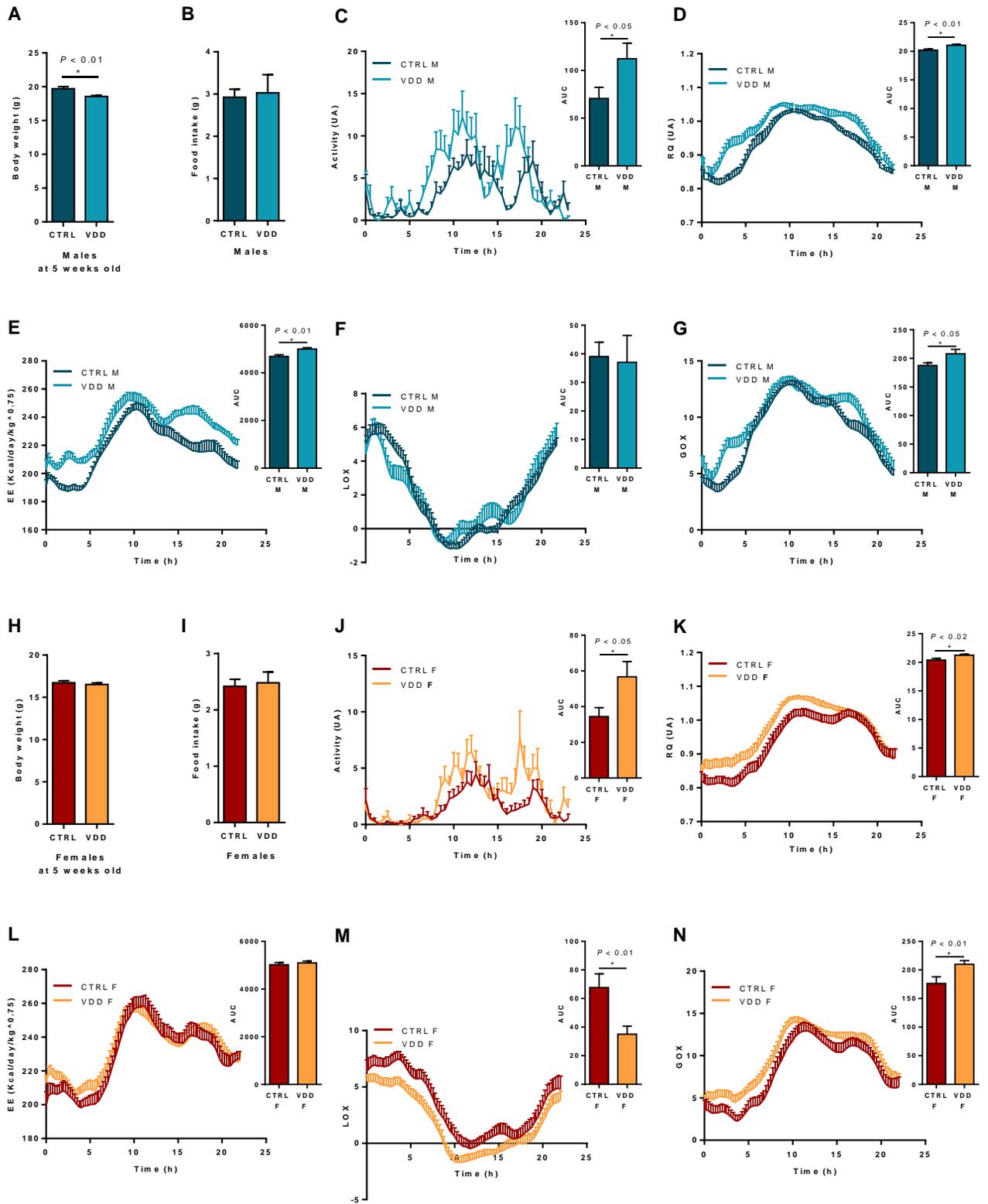


Fig. 2

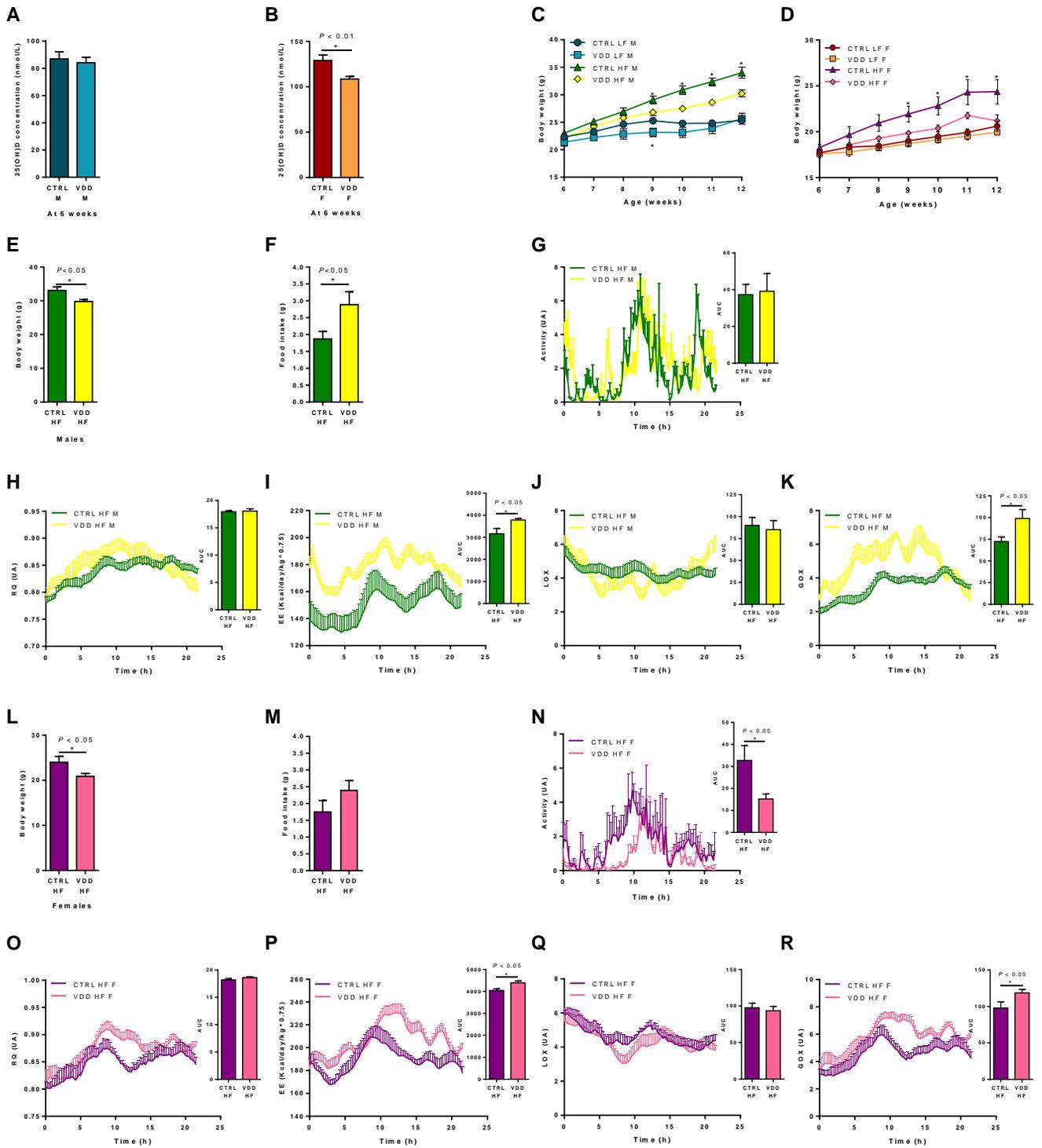


Fig. 3

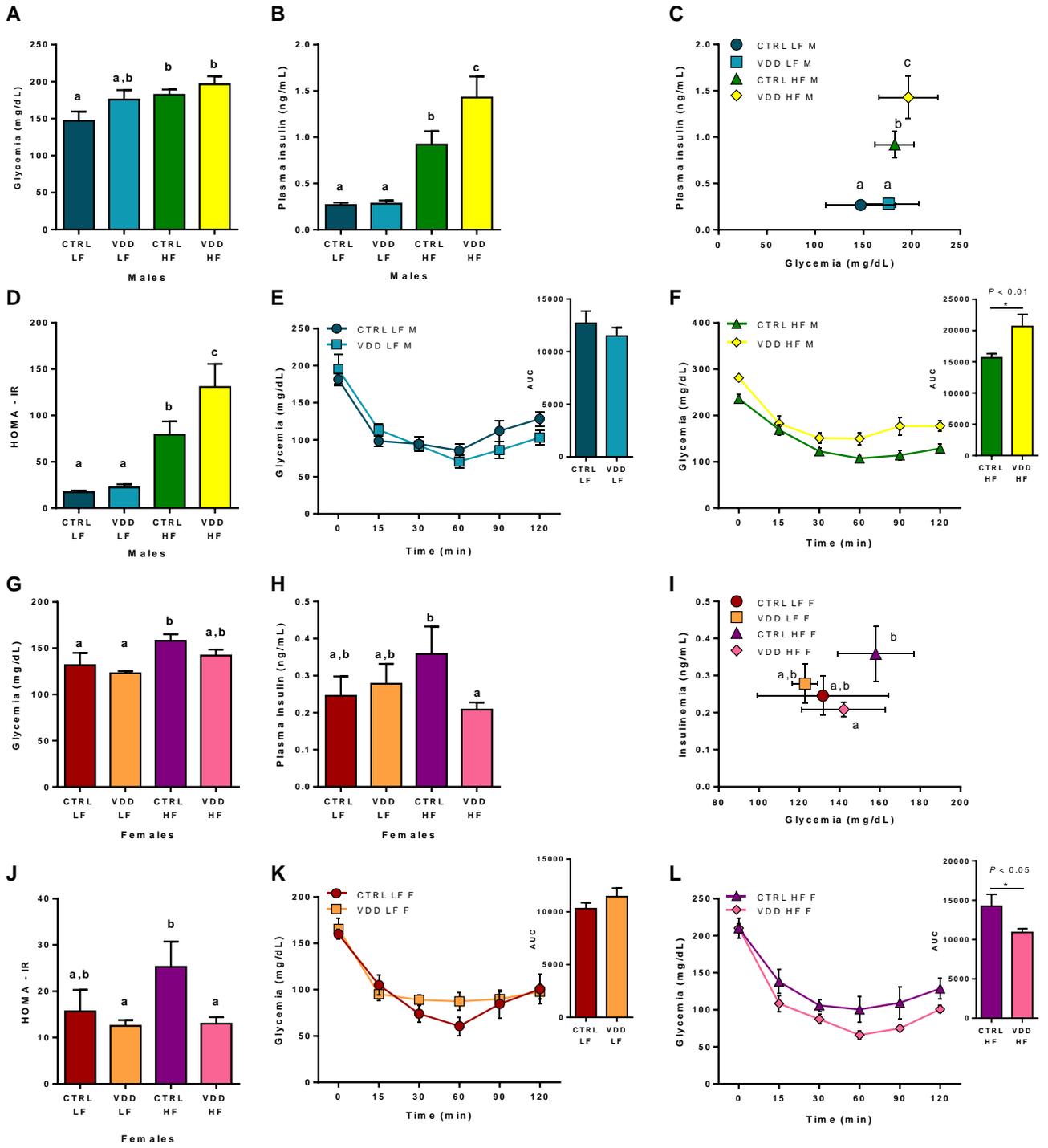


Fig. 4

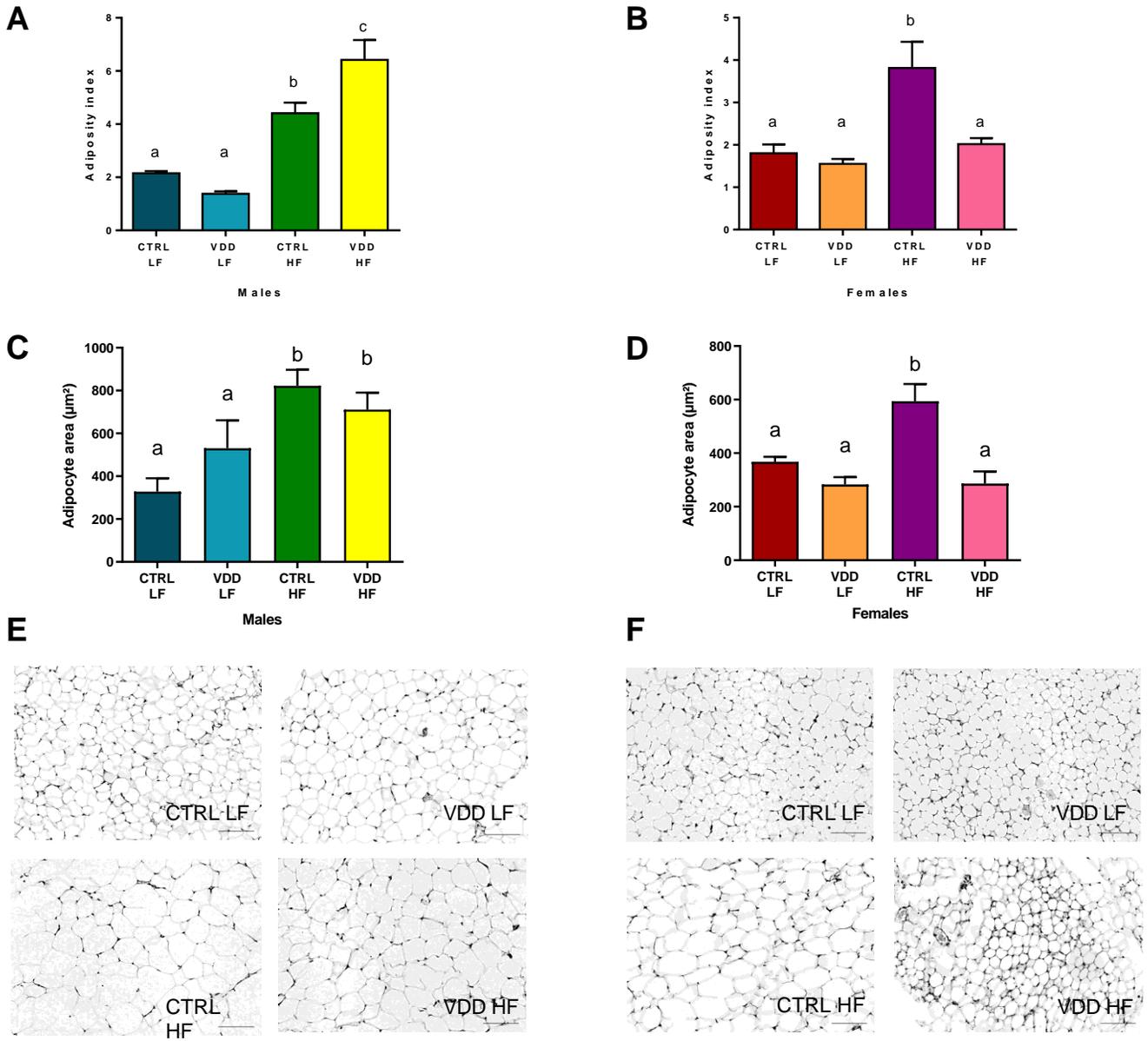
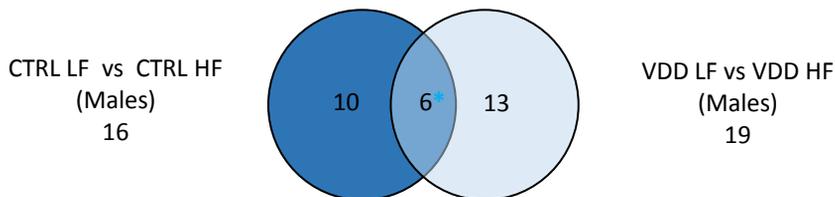
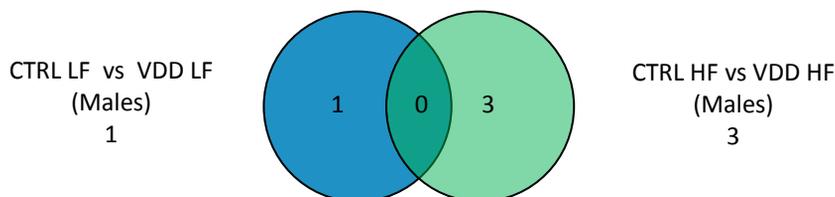


Fig. 5



Canonical Pathways	-log(p-value)	Z-score
GP6 Signaling Pathway*	6,67E00	2,982
Apelin Liver Signaling Pathway*	4,43E00	2,646
Apelin Cardiomyocyte Signaling Pathway	3,38E00	-0,302
Triacylglycerol Biosynthesis*	2,76E00	-2,646
Stearate Biosynthesis I (Animals)*	2,72E00	-1,890
γ-linolenate Biosynthesis II (Animals)	2,36E00	-2,000
Mitochondrial L-carnitine Shuttle Pathway	2,36E00	-1,000
LPS/IL-1 Mediated Inhibition of RXR Function	2,23E00	1,890
Valine Degradation I*	2,21E00	-2,000
CDP-diacylglycerol Biosynthesis I	2,07E00	-2,000
NRF2-mediated Oxidative Stress Response*	1,98E00	-1,342
Phosphatidylglycerol Biosynthesis II (Non-plastidic)	1,89E00	-2,000
Intrinsic Prothrombin Activation Pathway	1,84E00	2,000
Fatty Acid β-oxidation I	1,72E00	-2,236
Apelin Cardiac Fibroblast Signaling Pathway	1,55E00	1,000
p53 Signaling	1,38E00	-0,378

Canonical Pathways	-log(p-value)	Z-score
Oxidative Phosphorylation	6,92E00	-5,292
GP6 Signaling Pathway*	4,45E00	4,200
Triacylglycerol Biosynthesis*	4,22E00	-1,807
Nicotine Degradation II	2,56E00	-1,667
Valine Degradation I*	2,36E00	-2,449
Sirtuin Signaling Pathway	2,33E00	1,300
Apelin Liver Signaling Pathway*	2,28E00	1,890
NRF2-mediated Oxidative Stress Response*	1,93E00	-1,508
Stearate Biosynthesis I (Animals)*	1,78E00	-1,667
Superpathway of Melatonin Degradation	1,73E00	-0,707
Apelin Adipocyte Signaling Pathway	1,7E00	-1,941
Melatonin Degradation I	1,68E00	-1,134
Ethanol Degradation IV	1,64E00	-1,342
Nicotine Degradation III	1,52E00	-0,816
Isoleucine Degradation I	1,48E00	-2,000
Oxidative Ethanol Degradation III	1,48E00	-1,000
Ethanol Degradation II	1,48E00	-1,342
Glutathione-mediated Detoxification	1,4E00	-0,816
Bupropion Degradation	1,32E00	-2,000

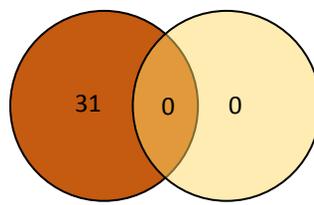


Canonical Pathways	-log(p-value)	Z-score
cAMP-mediated signaling	1,98E00	-1,000

Canonical Pathways	-log(p-value)	Z-score
Protein Kinase A Signaling	4,55E00	-0,447
Actin Cytoskeleton Signaling	4,32E00	2,236
RhoA Signaling	3,74E00	0,447

Fig. 6

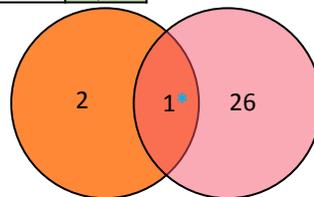
CTRL LF vs CTRL HF
(Females)
31



VDD LF vs VDD HF
(Females)
0

Canonical Pathways	-log(p-value)	Z-score
Valine Degradation I	6,19E00	-3,162
Fatty Acid β -oxidation I	4,62E00	-3,606
Oxidative Phosphorylation	4,44E00	-5,000
TCA Cycle II (Eukaryotic)	3,53E00	-3,162
Complement System	3,53E00	0,333
Branched-chain α -keto acid Dehydrogenase Complex	3,15E00	-2,000
Acetyl-CoA Biosynthesis I (Pyruvate Dehydrogenase Complex)	3,15E00	-2,000
Isoleucine Degradation I	2,99E00	-2,449
Sirtuin Signaling Pathway	2,75E00	1,372
Triacylglycerol Biosynthesis	2,62E00	-2,496
Apelin Adipocyte Signaling Pathway	2,53E00	-1,500
GP6 Signaling Pathway	2,2E00	2,837
Dendritic Cell Maturation	2,16E00	2,600
EIF2 Signaling	2,07E00	2,294
CDP-diacylglycerol Biosynthesis I	2	-1,414
LPS/IL-1 Mediated Inhibition of RXR Function	1,97E00	2,496
Calcium-induced T Lymphocyte Apoptosis	1,95E00	0,577
Glutaryl-CoA Degradation	1,94E00	-2,449
Salvage Pathways of Pyrimidine Ribonucleotides	1,94E00	-2,840
Glutathione Redox Reactions I	1,76E00	-1,633
Phosphatidylglycerol Biosynthesis II (Non-plastidic)	1,73E00	-1,414
Nicotine Degradation II*	1,73E00	-2,828
Mitochondrial L-carnitine Shuttle Pathway	1,65E00	-1,342
Intrinsic Prothrombin Activation Pathway*	1,6E00	1,342
Gluconeogenesis I*	1,6E00	-1,633
LXR/RXR Activation	1,47E00	-1,897
Ethanol Degradation IV	1,47E00	-2,236
Tryptophan Degradation III (Eukaryotic)	1,46E00	-2,449
Oxidative Ethanol Degradation III	1,35E00	-2,000
Coagulation System*	1,32E00	0,447
Ethanol Degradation II	1,32E00	-2,236

CTRL LF vs VDD LF
(Females)
3



CTRL HF vs VDD HF
(Females)
27

Canonical Pathways	-log(p-value)	Z-score
EIF2 Signaling	1,65E01	3,464
GP6 Signaling Pathway*	8,59E00	-2,530
Apelin Liver Signaling Pathway	4,8E00	-2,000

Canonical Pathways	-log(p-value)	Z-score
Oxidative Phosphorylation	4,77E01	7,211
Sirtuin Signaling Pathway	1,86E01	-3,212
TCA Cycle II (Eukaryotic)	1,61E01	4,000
Valine Degradation I	1,01E01	3,162
Fatty Acid β -oxidation I	8,11E00	3,464
Isoleucine Degradation I	5,25E00	2,449
Branched-chain α -keto acid Dehydrogenase Complex	4,73E00	2,000
Acetyl-CoA Biosynthesis I (Pyruvate Dehydrogenase Complex)	4,73E00	2,000
Ethanol Degradation II	4,68E00	2,646
Glutaryl-CoA Degradation	4,04E00	2,449
Tryptophan Degradation III (Eukaryotic)	3,44E00	2,449
Leucine Degradation I	3,25E00	2,000
Noradrenaline and Adrenaline Degradation	3,13E00	2,449
Oxidative Ethanol Degradation III	2,74E00	2,000
Intrinsic Prothrombin Activation Pathway	2,68E00	-1,000
Gluconeogenesis I	2,68E00	1,342
Mitochondrial L-carnitine Shuttle Pathway	2,37E00	2,000
LPS/IL-1 Mediated Inhibition of RXR Function	2,27E00	-1,890
Triacylglycerol Biosynthesis	2,23E00	2,646
Ethanol Degradation IV	2,22E00	2,000
Coagulation System	2,08E00	-1,000
Serotonin Degradation	2,06E00	2,449
GP6 Signaling Pathway*	2,06E00	-2,714
Nicotine Degradation II	1,9E00	1,342
Estrogen Biosynthesis	1,65E00	2,000
Stearate Biosynthesis I (Animals)	1,47E00	2,236
Superpathway of Methionine Degradation	1,41E00	2,000