

SUPPLEMENTARY FIGURES AND TABLES

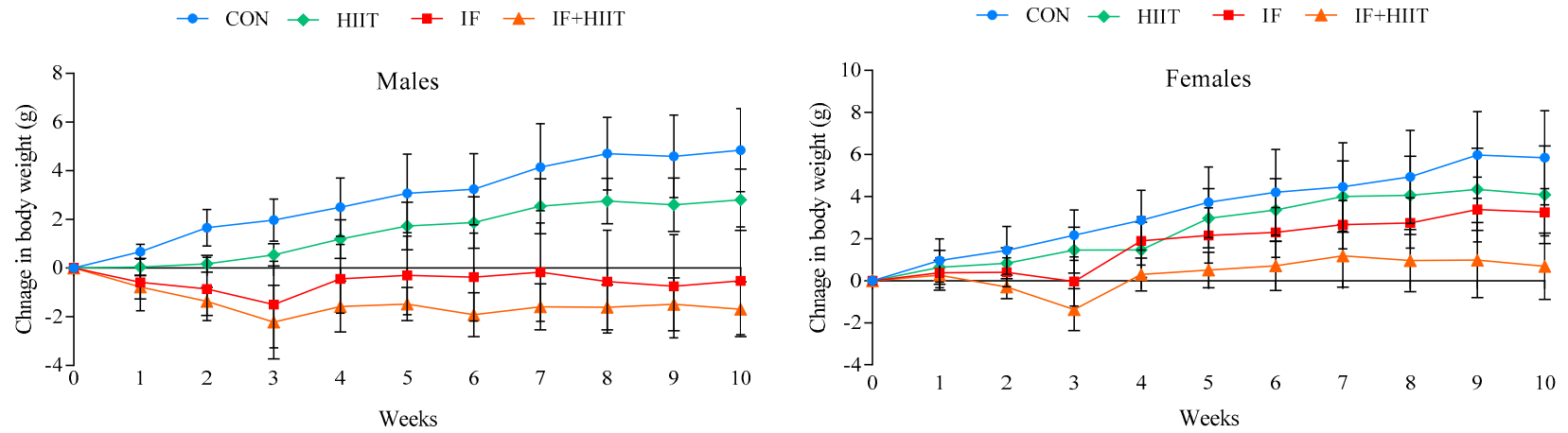


Figure S1. Change in body weight over the 10-weeks intervention period in (A) male and (B) females. Data is presented as mean of each group and error bars represent standard deviation.

Table S1. List of targeted genes in with full official names, NCBI reference sequence numbers and Qiagen catalogue number.

Gene symbol	Official full name	Marker	NCBI reference sequence number	Qiagen Catalogue number
<i>ACTB</i>	β -Actin, beta	House keeping	NM_007393	PPM02945
<i>AGRP</i>	Agouti related protein	Orexigenic	NM_007427	PPM29573
<i>AMPK</i>	Protein kinase AMP-activated catalytic subunit α 2	Energy sensor	NM_178143	PPM29410F
<i>AS160</i>	Akt substrate of 160 kDa	Insulin signalling	NM_00108127	PPM40317A
<i>B2M</i>	Beta-2 microglobulin	House keeping	NM_009735	PPM03562A
<i>BDNF</i>	Brain derived neurotrophic factor	Neuronal plasticity	NM_007540	PPM03006
<i>BiP</i>	Binding-Immunoglobulin Protein	Endoplasmic reticulum stress	NM_022310	PPM03586
<i>CIDEc</i>	Cell death-inducing DFFA-like effector c	Unilocular lipid droplet formation	NM_178373	PPM25558B
<i>COX-IV</i>	Cytochrome c oxidase subunit 4 isoform 2	Oxidative metabolism	NM_053091	PPM32878E
<i>CPT1</i>	Carnitine palmitoyltransferase Ib	Oxidative metabolism	NM_009948	PPM57688A
<i>CS</i>	Citrate synthase	Oxidative metabolism	NM_026444	PPM29621B
<i>FABP4</i>	Fatty acid binding protein 4, adipocyte	Fat mobilization	NM_024406	PPM04517A
<i>FOXO1</i>	Forkhead box O 1	Adipocyte differentiation and maturation	NM_019739	PPM03381C
<i>FTO</i>	Fat mass and obesity associated	Obesity	NM_011936	PPM39276
<i>GAPD</i>	Glyceraldehyde-3-phosphate dehydrogenase	House keeping	NM_008084	PPM02946
<i>HADH</i>	Hydroxyacyl-Coenzyme A dehydrogenase	β Oxidation	NM_008212	PPM35732B
<i>HIF1α</i>	Hypoxia inducible factor 1, alpha	Hypoxia	NM_010431	PPM03799C
<i>IL1β</i>	Interleukin 1 beta	Inflammation	NM_008361	PPM03109
<i>LEP</i>	Leptin	Obesity	NM_008493	PPM03504B
<i>MAFbx</i>	F-box protein 32	Muscle remodelling/atrophy	NM_026346	PPM38061A
<i>MuRF1</i>	Muscle RING finger 1	Muscle remodelling/atrophy	NM_00103904	PPM61645B
<i>NPY</i>	Neuropeptide Y	Orexigenic	NM_023456	PPM04323
<i>PGC1α</i>	Peroxisome proliferator-activated receptor gamma coactivator 1-alpha	Energy sensing & mitochondrial function	NR_027710	PPM03360I
<i>POMC</i>	Pro-opiomelanocortin-alpha	Anorexigenic	NM_008895	PPM37114
<i>PPARγ</i>	Peroxisome proliferator activated receptor γ	Adipocyte differentiation & maturation	NM_001127330	PPM05108C
<i>PPIA</i>	Peptidylprolyl isomerase A	House keeping	NM_008907	PPM03717B
<i>SIRT1</i>	Sirtuin 1	Energy sensor	NM_001159589	PPM05054A
<i>TNFα</i>	Tumor necrosis factor-alpha	Inflammation	NM_013693	PPM03113G

<i>TRPV4</i>	Transient receptor potential cation channel, subfamily V, member 4	Inflammation	NM_022017	PPM36070A
<i>UCP3</i>	Uncoupling protein 3	Fatty acid metabolism	NM_009464	PPM25342F
<i>XBP1us</i>	X-box binding protein 1 (unspliced)	Endoplasmic reticulum stress	NM_013842	PPM84308A
<i>XBP1s</i>	X-box binding protein 1 (spliced)	Endoplasmic reticulum stress	NM_001271730	PPM84307A

Table S2. List of miRNAs analysed in adipose tissue, miRbase accession numbers and Qiagen miScript Primer Assay Catalog Numbers.

Mature miRNA ID	Gene target	miRBase Accession No.	Qiagen miScript Primer Assay Catalog No.
mmu-miR-24-3p	<i>FABP4</i>	MIMAT0000219	MS00005922
mmu-miR-222-3p	<i>HIF1α</i>	MIMAT0000670	MS00007959
mmu-miR-143-3p	Leptin	MIMAT0000247	MS00001617
mmu-miR-145a-5p	<i>FOXO1</i>	MIMAT0000157	MS00001631
mmu-miR-696	<i>PGC1α</i>	MIMAT0003483	MS00002870
mmu-miR-133a-3p	<i>TNFα</i>	MIMAT0000145	MS00007294
mmu-miR-133b-3p	<i>MuRF1</i>	MIMAT0000769	MS00007301

Table S3. Pearson correlation between hypothalamus gene markers with body composition and glucose parameters

			AgRP	NPY	POMC	BDNF	TNF α	IL1B	BiP	FTO	XBPs/XBPus
Body Wt	Males	r	-0.190	0.093	0.091	-0.036	0.318	-0.261	0.321	-0.145	0.248
		<i>p value</i>	0.343	0.639	0.651	0.850	0.093	0.180	0.083	0.446	0.178
	Females	r	-0.188	0.402	-0.125	-0.021	0.147	-0.049	-0.096	-0.324	0.140
		<i>p value</i>	0.266	0.014	0.504	0.900	0.386	0.773	0.585	0.058	0.395
Fat Mass	Males	r	-0.090	0.091	0.087	0.014	0.250	-0.313	0.313	0.026	0.237
		<i>p value</i>	0.657	0.646	0.667	0.942	0.192	0.104	0.093	0.890	0.199
	Females	r	-0.198	0.422	-0.172	-0.032	0.108	-0.038	-0.092	-0.366	0.146
		<i>p value</i>	0.241	0.009	0.356	0.849	0.524	0.821	0.599	0.031	0.375
Lean mass	Males	r	-0.293	0.013	0.019	-0.139	0.241	0.032	0.116	-0.417	0.099
		<i>p value</i>	0.138	0.950	0.925	0.464	0.208	0.870	0.541	0.022	0.596
	Females	r	-0.057	0.160	0.149	0.081	0.300	-0.061	-0.100	-0.012	0.107
		<i>p value</i>	0.738	0.345	0.425	0.635	0.072	0.718	0.568	0.944	0.516
Fasting glucose	Males	r	-0.188	0.060	-0.053	-0.191	0.237	0.310	0.120	-0.385	0.037
		<i>p value</i>	0.348	0.762	0.792	0.312	0.217	0.108	0.527	0.036	0.843
	Females	r	-0.160	0.037	-0.068	-0.130	0.049	-0.011	0.130	-0.108	-0.008
		<i>p value</i>	0.344	0.828	0.718	0.444	0.775	0.946	0.455	0.537	0.961
Glucose AUC	Males	r	-0.170	0.426	-0.066	-0.227	0.202	0.145	-0.084	-0.317	-0.148
		<i>p value</i>	0.396	0.024	0.745	0.229	0.295	0.461	0.661	0.088	0.426
	Females	r	-0.236	0.093	-0.001	-0.161	-0.030	0.032	-0.053	-0.309	0.344
		<i>p value</i>	0.159	0.586	0.997	0.341	0.859	0.850	0.760	0.071	0.032
HOMA-IR	Males	r	-0.038	-0.060	0.091	-0.121	0.501	0.055	0.290	-0.055	0.292
		<i>p value</i>	0.851	0.763	0.651	0.525	0.006	0.780	0.120	0.773	0.111
	Females	r	-0.053	0.113	-0.107	0.032	0.050	-0.060	0.043	-0.075	-0.155
		<i>p value</i>	0.755	0.506	0.568	0.849	0.768	0.726	0.808	0.670	0.347

Table S4. Pearson correlation between adipose tissue gene markers with body composition, glucose parameters and plasma lipids.

			CIDEc	Leptin	PPAR	FOXO	HADH	FABP4	SIRT1	TRPV	HIF1a
Body Wt	Males	r	-0.157	0.722	-0.459	-0.069	-0.281	-0.596	-0.213	0.038	0.658
		<i>p value</i>	0.425	0.000	0.014	0.726	0.147	0.001	0.258	0.841	0.000
	Females	r	-0.005	0.671	-0.157	0.340	-0.126	-0.112	0.015	0.064	0.319
		<i>p value</i>	0.976	0.000	0.367	0.053	0.479	0.544	0.930	0.713	0.070
Fat Mass	Males	r	-0.107	0.785	-0.454	-0.008	-0.185	-0.574	-0.178	0.075	0.685
		<i>p value</i>	0.588	0.000	0.015	0.969	0.345	0.001	0.347	0.687	0.000
	Females	r	0.072	0.730	-0.129	0.366	-0.087	-0.063	0.058	0.106	0.349
		<i>p value</i>	0.692	0.000	0.462	0.036	0.625	0.733	0.735	0.541	0.047
Lean mass	Males	r	-0.179	0.107	-0.196	-0.148	-0.324	-0.258	-0.146	-0.051	0.149
		<i>p value</i>	0.363	0.576	0.317	0.451	0.093	0.177	0.443	0.786	0.458
	Females	r	-0.358	0.218	-0.226	0.159	-0.290	-0.295	-0.206	-0.149	0.075
		<i>p value</i>	0.041	0.208	0.192	0.378	0.096	0.101	0.222	0.387	0.677
Fasting glucose	Males	r	-0.289	0.098	-0.296	-0.153	-0.373	-0.417	-0.343	-0.114	0.137
		<i>p value</i>	0.136	0.605	0.126	0.437	0.051	0.024	0.064	0.542	0.495
	Females	r	-0.507	-0.111	-0.328	-0.151	-0.564	-0.412	-0.416	-0.300	0.082
		<i>p value</i>	0.003	0.526	0.055	0.401	0.001	0.019	0.010	0.076	0.650
Glucose AUC	Males	r	-0.030	0.511	-0.394	0.200	0.049	-0.416	-0.187	0.076	0.484
		<i>p value</i>	0.881	0.004	0.038	0.307	0.806	0.025	0.322	0.684	0.011
	Females	r	-0.274	0.279	-0.180	0.191	-0.177	-0.229	-0.264	-0.074	0.163
		<i>p value</i>	0.123	0.105	0.302	0.287	0.316	0.207	0.114	0.667	0.366
HOMA-IR	Males	r	-0.306	0.156	-0.160	-0.340	-0.451	-0.387	-0.076	-0.225	0.087
		<i>p value</i>	0.114	0.411	0.417	0.077	0.016	0.038	0.689	0.225	0.666
	Females	r	-0.283	0.089	-0.130	0.050	-0.384	-0.322	-0.375	-0.383	-0.080
		<i>p value</i>	0.111	0.611	0.455	0.783	0.025	0.073	0.022	0.021	0.657
TAG	Males	r	0.256	-0.121	0.118	0.246	-0.045	-0.127	-0.129	0.270	-0.069
		<i>p value</i>	0.198	0.532	0.560	0.206	0.819	0.512	0.504	0.149	0.734
	Females	r	0.126	0.102	-0.328	0.118	-0.029	0.320	0.229	0.074	0.029
		<i>p value</i>	0.487	0.558	0.054	0.514	0.870	0.074	0.173	0.667	0.872
HDL	Males	r	-0.182	0.067	-0.274	-0.056	-0.166	-0.216	-0.297	-0.112	0.063
		<i>p value</i>	0.355	0.726	0.159	0.779	0.398	0.261	0.111	0.549	0.756
	Females	r	0.132	0.424	-0.424	0.048	-0.016	-0.081	0.125	0.189	0.186
		<i>p value</i>	0.464	0.011	0.011	0.791	0.928	0.658	0.460	0.271	0.300
LDL	Males	r	0.029	0.600	-0.396	0.171	-0.077	-0.490	-0.089	0.254	0.679
		<i>p value</i>	0.882	0.001	0.037	0.385	0.696	0.007	0.639	0.168	0.000
	Females	r	0.169	0.364	0.148	0.157	0.201	0.029	0.122	0.187	0.272
		<i>p value</i>	0.346	0.032	0.397	0.383	0.255	0.873	0.473	0.274	0.126

Table S5. Pearson correlation between muscle gene markers with body composition and glucose parameters

			AMPK	SIRT1	PGC1	CPT1	CS	COX4	UCP3	TNFa	As160	MURF	MAFbx
Body Wt	Males	r	0.166	-0.129	0.491	-0.127	0.267	0.160	-0.131	0.502	-0.280	0.141	-0.154
		<i>p value</i>	0.400	0.505	0.006	0.510	0.161	0.408	0.514	0.007	0.149	0.485	0.434
	Females	r	0.019	0.044	0.304	0.267	0.031	-0.046	0.187	0.140	-0.199	-0.029	0.023
		<i>p value</i>	0.913	0.806	0.076	0.105	0.861	0.796	0.283	0.444	0.267	0.867	0.898
Fat Mass	Males	r	0.307	-0.005	0.492	-0.028	0.220	0.220	-0.008	0.498	-0.206	0.176	-0.029
		<i>p value</i>	0.112	0.978	0.006	0.887	0.252	0.251	0.968	0.007	0.294	0.379	0.883
	Females	r	0.029	0.111	0.342	0.282	0.046	-0.084	0.186	0.110	-0.171	-0.013	0.035
		<i>p value</i>	0.867	0.534	0.044	0.086	0.791	0.638	0.285	0.547	0.341	0.938	0.842
Lean mass	Males	r	-0.229	-0.294	0.163	-0.266	0.173	-0.099	-0.312	0.174	-0.232	-0.056	-0.326
		<i>p value</i>	0.240	0.122	0.390	0.163	0.369	0.609	0.113	0.377	0.235	0.782	0.091
	Females	r	-0.029	-0.296	0.017	0.093	-0.084	0.070	0.140	0.199	-0.237	-0.068	-0.024
		<i>p value</i>	0.863	0.089	0.925	0.578	0.633	0.696	0.423	0.275	0.184	0.692	0.890
Fasting glucose	Males	r	-0.119	-0.384	0.158	-0.403	-0.028	-0.109	-0.335	0.276	-0.425	-0.076	-0.284
		<i>p value</i>	0.546	0.040	0.406	0.030	0.883	0.575	0.087	0.155	0.024	0.707	0.143
	Females	r	-0.337	-0.350	-0.129	-0.234	-0.301	-0.074	-0.222	0.077	-0.373	-0.272	-0.254
		<i>p value</i>	0.042	0.042	0.459	0.157	0.080	0.676	0.200	0.677	0.033	0.108	0.141
Glucose AUC	Males	r	0.035	-0.234	0.164	-0.360	-0.036	-0.158	-0.203	0.249	-0.508	0.117	-0.094
		<i>p value</i>	0.859	0.221	0.388	0.055	0.852	0.413	0.311	0.201	0.006	0.561	0.635
	Females	r	-0.167	-0.143	-0.003	-0.110	-0.169	-0.097	-0.052	0.379	-0.366	0.032	0.072
		<i>p value</i>	0.323	0.418	0.987	0.511	0.332	0.585	0.768	0.032	0.036	0.851	0.681
HOMA-IR	Males	r	-0.079	-0.221	0.189	-0.174	0.049	0.012	-0.247	0.210	-0.246	-0.096	-0.204
		<i>p value</i>	0.690	0.249	0.319	0.367	0.799	0.952	0.214	0.283	0.207	0.633	0.297
	Females	r	-0.115	-0.284	-0.108	-0.070	-0.343	0.011	0.119	-0.206	-0.192	-0.079	-0.144
		<i>p value</i>	0.499	0.104	0.537	0.678	0.044	0.950	0.496	0.258	0.285	0.649	0.408

Table S6. Pearson correlation between mRNA and miRNA expression in adipose tissue

Gene	miRNA	Males		Females	
		r	P value	r	P value
FABP4	miR-24-3p	0.099	0.622	-0.069	0.713
HIF1	miR-222-3p	0.189	0.365	0.107	0.568
Leptin	miR-143-3p	-0.096	0.627	-0.295	0.096
FOXO1	miR-145-5p	0.476	0.016	-0.131	0.497

Table S7. Pearson correlation between mRNA and miRNA expression in muscle

Gene	miRNA	Males		Females	
		r	p value	r	p value
PGC1	miR-696	-0.065	0.752	0.488	0.009
TNF	miR-133a	-0.190	0.362	-0.404	0.041
MURF1	miR-133b	0.105	0.627	-0.270	0.164