

**Table S2:** Proteins and metabolites found differentially expressed in the comparison of INS-1E  $\beta$  cells treated with palmitate vs cells without treatment (grey lines when fold change  $\geq 1.5$ ).

<b>Fold palm/ctrl</b>	<b>ID</b>	<b>Name</b>	<b>Function</b>	<b>Location</b>
-1.48	P10860	glutamate dehydrogenase 1	aminoacid metabolism	mito
-1.21	Q05962	solute carrier family 25 (mitochondrial carrier; adenine nucleotide translocator), member 4	channels and transporter	mito
+1.43	P81155	voltage-dependent anion channel 2	channels and transporter	mito
+1.54	Q8K3P6	solute carrier family 25 (mitochondrial carrier; phosphate carrier), member 25	channels and transporter	mito
+1.61	P29410	adenylate kinase 2	kinase/phosphatases	mito
+2.22	Q9WUS0	adenylate kinase 4	kinase/phosphatases	mito
-1.69	P00507	glutamic-oxaloacetic transaminase 2	krebs cycle/ATP production	mito
-1.35	P21571	ATP synthase, H <sup>+</sup> transporting, mitochondrial Fo complex, subunit F6	krebs cycle/ATP production	mito
-1.23	P32551	ubiquinol-cytochrome c reductase core protein II	krebs cycle/ATP production	mito
-1.21	Q9ER34	aconitase 2, mitochondrial	krebs cycle/ATP production	mito
+1.28	P35435	ATP synthase, H <sup>+</sup> transporting, mitochondrial F1 complex, gamma polypeptide 1	krebs cycle/ATP production	mito
+1.30	P10719	ATP synthase, H <sup>+</sup> transporting, mitochondrial F1 complex, beta polypeptide	krebs cycle/ATP production	mito
+1.61	P13803	electron-transfer-flavoprotein, alpha polypeptide	krebs cycle/ATP production	mito
+1.70	P15999	ATP synthase, H <sup>+</sup> transporting, mitochondrial F1 complex, alpha subunit 1, cardiac muscle	krebs cycle/ATP production	mito
+2.00	P20788	ubiquinol-cytochrome c reductase, Rieske iron-sulfur polypeptide 1	krebs cycle/ATP production	mito
+2.27	P29419	ATP synthase, H <sup>+</sup> transporting, mitochondrial F1F0 complex, subunit E	krebs cycle/ATP production	mito
-1.15	HMDB00552	3-methylglutaryl carnitine	Lipid metabolism	mito
+1.04	HMDB13130	glutaryl carnitine	Lipid metabolism	mito
+1.25	P18886	carnitine palmitoyltransferase 2	Lipid metabolism	mito
+1.45	P15650	acyl-CoA dehydrogenase, long chain	Lipid metabolism	mito
+1.75	O35547	acyl-CoA synthetase long-chain family member 4	Lipid metabolism	mito
+1.96	P56720	sterol regulatory element binding transcription factor 1	Lipid metabolism	mito
-1.43	P07153	ribophorin I	metabolism/miscellaneous	mito
+1.11	Q561S0	NADH dehydrogenase (ubiquinone) 1 alpha subcomplex, 10,	mitochondrial enzymes	mito

		42kDa		
+1.47	P12007	isovaleryl-CoA dehydrogenase	mitochondrial enzymes	mito
+2.00	Q6AYQ8	fumarylacetoacetate hydrolase domain containing 1	mitochondrial enzymes	mito
+2.63	P07895	superoxide dismutase 2, mitochondrial	mitochondrial enzymes	mito
+1.18	Q6AXV4	SAMM50 sorting and assembly machinery component	mitochondrial protein processing	mito
+1.24	P85834	Tu translation elongation factor, mitochondrial	mitochondrial protein processing	mito
+1.28	Q4FZT0	stomatin (EPB72)-like 2	mitochondrial protein processing	mito
+1.43	P67779	prohibitin	mitochondrial protein processing	mito
+1.79	P56571	ES1 protein homolog, mitochondrial	other/unknown function	mito
-1.29	Q6TUG0	DnaJ (Hsp40) homolog, subfamily B, member 11	protein synthesis/translation regulation/protein folding/ER stress	mito
+1.43	P48721	heat shock 70kDa protein 9 (mortalin)	protein synthesis/translation regulation/protein folding/ER stress	mito
+1.64	P0C2N5	suppressor of variegation 4-20 homolog 1 (Drosophila)	epigenetic regulation	nucleus
+1.56	P84245	H3 histone, family 3A	other/unknown function	nucleus
+1.82	A9UMV8	H2A histone family, member J	other/unknown function	nucleus
+1.54	P17046	lysosomal-associated membrane protein 2	Proteasoma/Lysosoma/Autophagy	plasma membrane
+1.30	P59215	guanine nucleotide binding protein (G protein), alpha activating activity polypeptide O	signal transduction	plasma membrane
+1.30	P04897	guanine nucleotide binding protein (G protein), alpha inhibiting activity polypeptide 2	signal transduction	plasma membrane
+1.37	P69897	tubulin, beta class I	cytoskeleton related proteins	cyto
+1.49	P47819	glial fibrillary acidic protein	cytoskeleton related proteins	cyto
+1.56	P68370	tubulin, alpha 1a	cytoskeleton related proteins	cyto
+1.69	P60711	actin, beta	cytoskeleton related proteins	cyto
+1.32	P04764	enolase 1, (alpha)	glucose metabolism	cyto
+1.49	P04797	glyceraldehyde-3-phosphate dehydrogenase	glucose metabolism	cyto
+2.00	Q9ESV6	glyceraldehyde-3-phosphate dehydrogenase, spermatogenic	glucose metabolism	cyto
-1.65	Q5RJR8	leucine rich repeat containing 59	other/unknown function	cyto
+2.08	Q63716	peroxiredoxin 1	oxidative stress/apoptosis	cyto
+1.67	P62986	ubiquitin A-52 residue ribosomal protein fusion product 1	posttranslational modification/ubiquitination	cyto
-1.24	P34058	heat shock protein 90kDa alpha (cytosolic), class B member 1	protein synthesis/translation regulation/protein folding/ER stress	cyto
+1.25	Q07984	signal sequence receptor, delta	protein synthesis/translation regulation/protein folding/ER stress	cyto
+1.28	P62630	eukaryotic translation elongation factor 1 alpha 1	protein synthesis/translation regulation/protein	cyto

+1.37	P19944	ribosomal protein, large, P1	folding/ER stress protein synthesis/translation regulation/protein folding/ER stress	cyto
+1.41	P62890	ribosomal protein L30	protein synthesis/translation regulation/protein folding/ER stress	cyto
+1.29	P38406	guanine nucleotide binding protein (G protein), alpha activating activity polypeptide, olfactory type	signal transduction	cyto
+1.40	P10536	RAB1B, member RAS oncogene family	vesicle transport/signal transduction	cyto
+1.47	Q63942	RAB3D, member RAS oncogene family	vesicle transport/signal transduction	cyto
+1.40	Q9WVB1	RAB6A, member RAS oncogene family	vesicle transport/signal transduction	cyto
-1.28	Q9JK11	reticulon 4	other/unknown function	ER
+1.67	P61621	Sec61 alpha 1 subunit ( <i>S. cerevisiae</i> )	other/unknown function	ER
-1.26	P85007	Immediate early response 3-interacting protein 1	oxidative stress/apoptosis	ER
-1.24	P04785	prolyl 4-hydroxylase, beta polypeptide	posttranslational modification/ubiquitination	ER
-1.41	Q63617	hypoxia up-regulated 1	protein synthesis/translation regulation/protein folding/ER stress	ER
-1.40	P18418	calreticulin	protein synthesis/translation regulation/protein folding/ER stress	ER
-1.28	P38659	protein disulfide isomerase family A, member 4	protein synthesis/translation regulation/protein folding/ER stress	ER
+1.24	P28576	platelet-derived growth factor alpha polypeptide	hormones/growth factors/receptors/neuropeptides	extracellular space
+1.29	P01323	insulin	hormones/growth factors/receptors/neuropeptides	extracellular space
+2.13	P01322	insulin I	hormones/growth factors/receptors/neuropeptides	extracellular space
+2.08	P08426	RIKEN cDNA 2210010C04 gene	other/unknown function	extracellular Space
-1.41	HMDB00182	L-lysine	aminoacid metabolism	other
-1.38	HMDB00158	L-tyrosine	aminoacid metabolism	other
-1.37	HMDB00687	L-leucine	aminoacid metabolism	other
-1.33	HMDB00517	L-arginine	aminoacid metabolism	other
-1.28	HMDB00148	L-glutamic acid	aminoacid metabolism	other
-1.28	HMDB00159	L-phenylalanine	aminoacid metabolism	other
-1.27	HMDB00168	L-asparagine	aminoacid metabolism	other
-1.12	HMDB00162	L-proline	aminoacid metabolism	other
-1.11	HMDB00214	L-ornithine	aminoacid metabolism	other

-1.09	HMDB00161	L-alanine	aminoacid metabolism	other
-1.09	HMDB00883	L-valine	aminoacid metabolism	other
+1.06	HMDB00177	L-histidine	aminoacid metabolism	other
+1.09	541-15-1	L-carnitine	aminoacid metabolism	other
+1.11	HMDB00187	L-serine	aminoacid metabolism	other
+1.16	HMDB00696	L-methionine	aminoacid metabolism	other
+1.17	HMDB00191	L-aspartic acid	aminoacid metabolism	other
+1.39	HMDB00167	L-threonine	aminoacid metabolism	other
+1.43	HMDB00123	glycine	aminoacid metabolism	other
+1.25	Q64119	myosin, light polypeptide 6, alkali, smooth muscle and non-muscle-like	other/unknown function	other
-1.21	P63018	heat shock 70kDa protein 8	protein synthesis/translation regulation/protein folding/ER stress	other

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