

| Gene Symbol | Gene Title | Entrez Gene | Leydig 3 week log2(KO / WT) | Leydig adult log2(KO / WT) | Gene Symbol | Gene Title | Entrez Gene | Leydig 3 week log2(KO / WT) | Leydig adult log2(KO / WT) |
|---------------|--|-------------|-----------------------------|----------------------------|-------------|---|-------------|-----------------------------|----------------------------|
| O610007P14Rik | RIKEN cDNA O610007P14 gene | 58520 | -0.266 | 0.367 | Myd88 | myeloid differentiation primary respor | 17874 | -0.473 | -0.414 |
| Abcb1a | ATP-binding cassette, sub-family B (MI | 18671 | -0.506 | -0.399 | Myoc | myocilin | 17926 | -0.562 | 0.105 |
| Acdb3 | acyl-Coenzyme A binding domain cont | 170760 | 0.120 | 0.246 | Ncoa1 | nuclear receptor coactivator 1 | 17977 | -0.283 | -0.496 |
| Adh1 | alcohol dehydrogenase 1 (class I) | 11522 | -0.428 | -0.831 | Ncoa2 | nuclear receptor coactivator 2 | 17978 | -0.715 | -0.649 |
| Calcoco1 | calcium binding and coiled coil domain | 67488 | -0.675 | -0.296 | Ncoa3 | nuclear receptor coactivator 3 | 17979 | -0.346 | 0.038 |
| Carrm1 | coactivator-associated arginine methyl | 59035 | -0.161 | 0.156 | Nenf | neuron derived neurotrophic factor | 66208 | -0.279 | 0.666 |
| Ccdc80 | coiled-coil domain containing 80 | 67896 | 0.899 | 0.446 | Npc11 | NPC1-like 1 | 237636 | 0.475 | 0.858 |
| Cebpb | CCAAT/enhancer binding protein (C/E | 12608 | 0.470 | 1.349 | Nphs2 | nephrosis 2 homolog, podocin (human | 170484 | -0.072 | -0.570 |
| Cited1 | Cbp/p300-interacting transactivator w | 12705 | -0.314 | -0.513 | Nr1d2 | nuclear receptor subfamily 1, group D, | 353187 | -1.097 | -0.937 |
| Cubn | cubilin (intrinsic factor-cobalamin rece | 65969 | 0.023 | 0.520 | Nr1h2 | nuclear receptor subfamily 1, group H, | 22260 | -0.149 | 0.397 |
| Cyb5 | cytochrome b-5 | 109672 | -0.663 | -0.413 | Nr1h3 | nuclear receptor subfamily 1, group H, | 22259 | -0.022 | 0.392 |
| Cyb5b | cytochrome b5 type B | 66427 | -0.291 | -0.196 | Nr2f2 | nuclear receptor subfamily 2, group F, | 11819 | -0.538 | 0.513 |
| Cyb5d2 | cytochrome b5 domain containing 2 | 192986 | -0.703 | -0.816 | Nr3c1 | nuclear receptor subfamily 3, group C, | 14815 | -0.109 | -0.376 |
| Cyb5r1 | cytochrome b5 reductase 1 | 72017 | -0.310 | -0.377 | Nr4a1 | nuclear receptor subfamily 4, group A, | 15370 | 0.435 | 0.224 |
| Cyb5r2 | cytochrome b5 reductase 2 | 320635 | 0.452 | -0.036 | Nr4a2 | Nuclear receptor subfamily 4, group A, | 18227 | 0.877 | -0.654 |
| Cyb5r3 | cytochrome b5 reductase 3 | 109754 | 0.143 | 0.073 | Nsdhl | NAD(P) dependent steroid dehydroge | 18194 | -0.259 | -0.294 |
| Cyb5r4 | cytochrome b5 reductase 4 | 266690 | 0.313 | 0.064 | Osbp | oxysterol binding protein | 76303 | -0.660 | -0.013 |
| Cyp11a1 | cytochrome P450, family 11, subfamily | 13070 | 0.023 | -0.031 | Osbp2 | oxysterol binding protein 2 | 74309 | 0.744 | 0.562 |
| Cyp17a1 | cytochrome P450, family 17, subfamily | 13074 | 0.012 | -0.241 | Osbp11 | oxysterol binding protein-like 11 | 106326 | -0.523 | -0.539 |
| Cyp24a1 | cytochrome P450, family 24, subfamily | 13081 | 0.277 | -0.594 | Osbp1a | oxysterol binding protein-like 1A | 64291 | -0.065 | 0.020 |
| Cyp39a1 | cytochrome P450, family 39, subfamily | 56050 | 0.053 | 3.064 | Osbp12 | oxysterol binding protein-like 2 | 228983 | 0.388 | -0.127 |
| Cyp3a13 | cytochrome P450, family 3, subfamily | 13113 | 0.905 | 0.804 | Paqr5 | progesterin and adipoQ receptor family | 74090 | 0.868 | 0.682 |
| Cyp46a1 | cytochrome P450, family 46, subfamily | 13116 | -0.548 | 0.029 | Paqr7 | Progesterin and adipoQ receptor family | 71904 | 0.522 | 1.041 |
| Cyp51 | cytochrome P450, family 51 | 13121 | 0.416 | 0.112 | Paqr8 | progesterin and adipoQ receptor family | 74229 | 0.051 | -0.215 |
| Dhcr24 | 24-dehydrocholesterol reductase | 74754 | 0.697 | 0.132 | Pgrmc1 | progesterone receptor membrane con | 53328 | 0.022 | -0.261 |
| Dhcr7 | 7-dehydrocholesterol reductase | 13360 | -0.331 | -0.955 | Pgrmc2 | progesterone receptor membrane con | 70804 | -0.397 | -0.049 |
| Dyrk1a | dual-specificity tyrosine-(Y)-phosphory | 13548 | -1.126 | -0.736 | Pmkv | phosphomevalonate kinase | 68603 | 0.894 | 0.224 |
| Ebp | phenylalkylamine Ca2+ antagonist (em | 13595 | -0.151 | -0.209 | Prkar1a | protein kinase, cAMP dependent regul | 19084 | 0.247 | -0.009 |
| Emp2 | epithelial membrane protein 2 | 13731 | 0.478 | 0.399 | Rhoa | ras homolog gene family, member A | 11848 | -0.083 | 0.276 |
| Esr1 | estrogen related receptor, alpha | 26379 | -0.114 | -0.221 | Rora | RAR-related orphan receptor alpha | 19883 | 1.176 | 0.880 |
| Esr2 | estrogen related receptor, beta | 26380 | -1.118 | 0.610 | Runx2 | runt related transcription factor 2 | 12393 | 0.693 | 0.456 |
| Fa2h | fatty acid 2-hydroxylase | 338521 | 0.239 | 0.223 | Rxr1 | retinoid X receptor alpha | 20181 | 0.578 | 0.509 |
| Fads1 | fatty acid desaturase 1 | 76267 | -0.036 | -0.602 | Rxr2 | retinoid X receptor beta | 20182 | -0.876 | 0.420 |
| Fads2 | fatty acid desaturase 2 | 56473 | -0.094 | -0.468 | Sc4mol | sterol-C4-methyl oxidase-like | 66234 | -0.607 | -0.961 |
| Fdft1 | farnesyl diphosphate farnesyl transfer | 14137 | 0.243 | 0.007 | Sc5d | sterol-C5-desaturase (fungal ERG3, del | 235293 | -0.434 | -0.599 |
| Fdps | farnesyl diphosphate synthetase | 110196 | 0.498 | -0.042 | Scap | SREBF chaperone | 235623 | -0.686 | -0.435 |
| Fdxr | ferredoxin reductase | 14149 | 0.180 | -0.363 | Scarb1 | scavenger receptor class B, member 1 | 20778 | 0.261 | 0.147 |
| Fkbp4 | FK506 binding protein 4 | 14228 | 0.001 | 0.177 | Sf1 | splicing factor 1 | 22668 | -0.132 | -0.233 |
| Gsk3b | glycogen synthase kinase 3 beta | 56637 | 1.195 | 0.798 | Slc37a4 | solute carrier family 37 (glucose-6-pho | 14385 | 0.867 | 0.388 |
| H2-Ke6 | H2-K region expressed gene 6 | 14979 | 0.255 | -0.162 | Smad2 | MAD homolog 2 (Drosophila) | 17126 | -0.332 | -0.053 |
| Hdlbp | high density lipoprotein (HDL) binding | 110611 | -0.383 | -0.257 | Smad3 | MAD homolog 3 (Drosophila) | 17127 | -0.361 | -1.054 |
| Hmgcr | 3-hydroxy-3-methylglutaryl-Coenzyme | 15357 | 0.223 | -0.278 | Smarca2 | SWI/SNF related, matrix associated, ac | 67155 | -0.462 | -0.010 |
| Hmgcs2 | 3-hydroxy-3-methylglutaryl-Coenzyme | 15360 | 0.208 | -0.409 | Smarca4 | SWI/SNF related, matrix associated, ac | 20586 | -0.675 | -0.190 |
| Hsd17b11 | hydroxysteroid (17-beta) dehydrogena | 114664 | -0.348 | -0.300 | Snai1 | snail homolog 1 (Drosophila) | 20613 | -0.040 | -0.182 |
| Hsd17b12 | hydroxysteroid (17-beta) dehydrogena | 56348 | -0.177 | -0.519 | Soat1 | sterol O-acyltransferase 1 | 20652 | -0.199 | -0.724 |
| Hsd17b2 | hydroxysteroid (17-beta) dehydrogena | 15486 | 0.373 | -0.171 | Sor11 | sortilin-related receptor, LDLR class A | 20660 | -0.067 | -0.284 |
| Hsd17b3 | hydroxysteroid (17-beta) dehydrogena | 15487 | -0.052 | -0.554 | Sqle | squalene epoxidase | 20775 | 0.071 | 0.014 |
| Hsd17b7 | hydroxysteroid (17-beta) dehydrogena | 15490 | 0.337 | -0.655 | Sra1 | sterol receptor RNA activator 1 | 24068 | -0.053 | -0.123 |
| Hsd3b1 | hydroxy-delta-5-steroid dehydrogenas | 15492 | -0.493 | -0.914 | Srd5a1 | steroid 5 alpha-reductase 1 | 78925 | -0.476 | 0.384 |
| Hsd3b7 | hydroxy-delta-5-steroid dehydrogenas | 101502 | 0.028 | -0.083 | Sreb1 | sterol regulatory element binding tran | 20787 | 0.278 | -0.443 |
| Idi1 | isopentenyl-diphosphate delta isomer | 319554 | -0.594 | -0.463 | Sreb2 | sterol regulatory element binding fact | 20788 | 0.273 | 0.177 |
| Igf1 | insulin-like growth factor 1 | 16000 | -0.082 | -0.412 | Star | steroidogenic acute regulatory protein | 20845 | -0.308 | -0.745 |
| Insig1 | insulin induced gene 1 | 231070 | -0.493 | -0.480 | Stard5 | STAR-related lipid transfer (START) don | 170460 | 0.754 | -0.601 |
| Insig2 | insulin induced gene 2 | 72999 | -0.258 | -0.365 | Stard6 | STAR-related lipid transfer (START) don | 170461 | 0.330 | 0.246 |
| Kl | klotho | 16591 | -0.693 | 0.898 | Stat5a | signal transducer and activator of tran | 20850 | -0.397 | -0.730 |
| Ldlr | low density lipoprotein receptor | 16835 | -0.262 | -0.496 | Sult1a1 | sulfotransferase family 1A, phenol-pre | 20887 | -0.701 | 1.075 |
| Lhb | luteinizing hormone beta | 16866 | -0.025 | 0.337 | Sult1e1 | sulfotransferase family 1E, member 1 | 20860 | -0.645 | -1.397 |
| Lhcgr | luteinizing hormone/choriogonadotro | 16867 | -0.915 | -1.107 | Sult4a1 | sulfotransferase family 4A, member 1 | 29859 | 0.940 | 0.955 |
| Lipa | lysosomal acid lipase A | 16889 | 0.782 | -0.654 | Suox | sulfite oxidase | 211389 | -0.303 | 0.160 |
| Lipe | lipase, hormone sensitive | 16890 | 0.672 | 0.453 | Tbl1x | transducin (beta)-like 1 X-linked | 21372 | 0.802 | 1.617 |
| Lss | lanosterol synthase | 16987 | -0.315 | 0.074 | Thrb | thyroid hormone receptor beta | 21834 | -1.733 | 0.787 |
| Mbtps1 | membrane-bound transcription factor | 56453 | 0.025 | -0.253 | Tm7sf2 | transmembrane 7 superfamily membe | 73166 | -0.004 | -0.141 |
| Mbtps2 | membrane-bound transcription factor | 270669 | -0.304 | -0.128 | Wwox | WW domain-containing oxidoreductas | 80707 | -0.499 | -0.529 |
| Mvd | mevalonate (diphospho) decarboxylas | 192156 | 0.320 | 0.373 | | | | | |
| Mvk | mevalonate kinase | 17855 | 0.097 | 0.159 | | | | | |

Supplemental Table 1 - Steroid and cholesterol biosynthesis genes represented in Figure 3B.

| Gene Symbol | Gene Title | 3 wk WT | adult WT | 3 wk KO | adult KO |
|-------------|---|-----------|----------|-----------|-----------|
| GLG1 | golgi apparatus protein 1 | 0.114008 | 1.30649 | -1.078474 | -0.342023 |
| BMP6 | bone morphogenetic protein 6 | -1.046307 | 1.329982 | -0.361103 | 0.077428 |
| CLCN3 | chloride channel 3 | -0.442321 | 1.265443 | -1.068919 | 0.245797 |
| NRP1 | neuropilin 1 | -0.654155 | 1.342431 | -0.860422 | 0.172146 |
| PPP3CB | protein phosphatase 3, catalytic subunit, beta isoform | -0.595563 | 1.33366 | -0.914146 | 0.176049 |
| CTNNB1 | catenin (cadherin associated protein), beta 1 | -0.752331 | 1.253985 | -0.85733 | 0.355676 |
| YWHAB | tyrosine 3-monooxygenase/tryptophan 5-monooxygenase activation protein, beta polypeptide | -0.775892 | 1.166821 | -0.888324 | 0.497395 |
| GNA13 | guanine nucleotide binding protein, alpha 13 | 1.075171 | 0.460277 | -0.291619 | -1.243829 |
| HDGF | Hepatoma-derived growth factor | 0.978877 | 0.614725 | -0.352881 | -1.240721 |
| ADCY9 | adenylate cyclase 9 | 0.820709 | 0.815178 | -0.41554 | -1.220347 |
| MYLK | myosin, light polypeptide kinase | 0.675094 | 0.967448 | -0.461188 | -1.181354 |
| PPP1R8 | protein phosphatase 1, regulatory (inhibitor) subunit 8 | 0.74519 | 0.905854 | -0.464174 | -1.186871 |
| KCTD12 | potassium channel tetramerisation domain containing 12 | 1.000527 | 0.684571 | -0.607463 | -1.077635 |
| MAP2K5 | | 0.872807 | 0.833638 | -0.644323 | -1.062122 |
| AKAP11 | | 0.331405 | 1.268488 | -0.825267 | -0.774625 |
| AKAP9 | A kinase (PRKA) anchor protein (yotiao) 9 | 0.405346 | 1.226483 | -0.815262 | -0.816567 |
| CREBBP | | 0.592881 | 1.078594 | -1.045752 | -0.625723 |
| YWHAZ | tyrosine 3-monooxygenase/tryptophan 5-monooxygenase activation protein, zeta polypeptide | 0.537288 | 1.130454 | -0.980013 | -0.687728 |
| CLCN6 | | 0.672539 | 1.037928 | -0.793804 | -0.916664 |
| ITPR1 | inositol 1,4,5-triphosphate receptor 1 | 0.762897 | 0.963002 | -0.837937 | -0.887963 |
| PYGB | brain glycogen phosphorylase | 0.93245 | 0.79561 | -0.911184 | -0.816876 |
| SMAD4 | MAD homolog 4 (Drosophila) | 0.886398 | 0.840993 | -0.950558 | -0.776832 |
| CHUK | conserved helix-loop-helix ubiquitous kinase | 1.18771 | 0.466799 | -0.864296 | -0.790213 |
| YWHAG | tyrosine 3-monooxygenase/tryptophan 5-monooxygenase activation protein, gamma polypeptide | 1.079405 | 0.592444 | -1.043716 | -0.628133 |
| H1F0 | H1 histone family, member 0 | 1.12136 | 0.194218 | -0.009924 | -1.305654 |
| CLCN2 | chloride channel 2 | 1.22239 | 0.381336 | -0.609018 | -0.994708 |
| CREB1 | | 1.199677 | 0.415056 | -0.601428 | -1.013305 |
| KCTD3 | | 1.230501 | 0.339904 | -0.523242 | -1.047163 |
| PIK3R4 | phosphatidylinositol 3 kinase, regulatory subunit, polypeptide 4, p150 | 1.14204 | 0.528808 | -0.73482 | -0.936027 |
| PPP1R12A | | 1.144663 | 0.512395 | -0.663789 | -0.993268 |
| GRB2 | growth factor receptor bound protein 2 | 1.31958 | 0.050069 | -0.28593 | -1.083719 |
| ATF1 | activating transcription factor 1 | 1.262101 | 0.189935 | -0.328008 | -1.124027 |
| PRKAR1A | protein kinase, cAMP dependent regulatory, type I, alpha | 1.288619 | 0.122023 | -0.299355 | -1.111287 |
| PLCG1 | | 1.322969 | 0.10884 | -0.389663 | -1.042146 |
| PRKD3 | protein kinase D3 | 1.355961 | 0.029377 | -0.375711 | -1.009627 |
| ADD3 | adducin 3 (gamma) | 1.319746 | 0.156408 | -0.469477 | -1.006677 |
| POLR2B | polymerase (RNA) II (DNA directed) polypeptide B | 1.343698 | 0.125439 | -0.511455 | -0.957682 |
| PDE3B | | 1.306648 | 0.222507 | -0.572221 | -0.956934 |

Supplemental Table 2 – cAMP-, PKA-, and CREB-regulated genes in Leydig cells represented in Figure 3H.

| | 3wk WT vs KO | adult WT vs KO |
|-------|--------------|----------------|
| Nr5a1 | -1.41 | 1 |
| Gata4 | -1.15 | 1.15 |
| Gata6 | -1.15 | -1.11 |
| Crem | -1.11 | 1.07 |
| Yy1 | 1.15 | 1.07 |
| Creb1 | 1 | 1.07 |
| Creb3 | 1 | -1.07 |
| Ap1s1 | 1.07 | 1.07 |
| Ap1b1 | 1.07 | -1.15 |

testes

| | 3wk WT vs KO | adult WT vs KO |
|-------|--------------|----------------|
| Nr5a1 | -1.09 | 1.14 |
| Gata4 | 1.19 | 1.29 |
| Gata6 | 1 | -1.37 |
| Crem | -1.17 | -1.19 |
| Yy1 | 1.66 | 1.49 |
| Creb1 | -1.03 | 1 |
| Creb3 | 1 | 1.16 |
| Ap1s1 | 1.16 | 1.05 |
| Ap1b1 | 1.24 | 1.13 |
| Ap1g1 | -1.33 | -1.01 |

Leydig cells

Supplemental Table 3 – Transcription factors targeting meiotic process genes are mostly unchanged in testes and Leydig cells of *Mrp4* KO mice. Selected transcription factors that target various meiotic processes have similar expression patterns in both testes and Leydig cells. Data are expressed as fold difference in *Mrp4*^{+/+} vs. *Mrp4*^{-/-} mice, calculated as the log₂ difference in the geometric mean of the signals. Positive values were raised to the power 2. If the log difference was negative, values were the negative reciprocal of the exponential value.

| | Forward Primer | Reverse Primer |
|---------|----------------------------|------------------------------|
| Lhr | GCGCGCCGGATGGT | AAAGCTTGTGATGGGATTACTTTGA |
| Star | CGGGTGGATGGGTCAAGTT | GGACAGCTCCTGGTCACTATAGAGT |
| Pbr | CTGGACACTGGCTCCCATCT | CCAGCTCTTTCCAGACTATGTAGGA |
| Cyp11a1 | GGCCGGCGGATTGC | CATCACGGAGATTTTGAACITCAA |
| Hsd3b1 | CCCAGGCAGACCATCCTAGA | GAACACAGGCCTCCAATAGGTT |
| Cyp17a1 | CCAGAGAAGTGCTCGTGAAGAA | CCTTTTCCTTGGTCCGACAA |
| Cyp2b9 | TGCTCAAGTACCCCATGTCA | AGAGAAAGTCCAAAAGGAGA |
| Cyp2b10 | TCTTCCAACGTTCCCCATTG | TCTTCCAACGTTCCCCATTG |
| Por | TCTTCCAACGTTCCCCATTG | TGCTCCCTCCATGTGATGAA |
| Cyp7a1 | AATGCACTTGGATCCTGAAATCTAC | TTCATCAAGGTACCGGTCGTATT |
| Cyp7b1 | CGGCCCTGTTCTCCTTAC | GATAAGGAAGCCAACCTTTTATCAAG |
| Cyp3a11 | CCAAACCTCTGCCATTTTTAGG | GCACTCCATGTGCAATTTCCA |
| Cyp3a13 | GCTGATGATAAACTCCAGAATTACAA | CCAAAGAATCTCATAAAGC |
| Cyp1a2 | ATCCTTTGTCCCCTTACCAT | GGGAATGTGGAAGCCATTCA |
| Gapdh | GTCCCGTAGACAAAATGGTGAAG | TCGGTGTGAACGGATT |
| AR | TGTCGTCTCCGGAATGTTATG | GTTTTAGATTTCCAAGTTTCTTCAGCTT |
| Actin | AGCCATGTACGTAGCCATCC | CTCTCAGCTGTGGTGGTGAA |
| Slco6b1 | TCGGAGGAAGAGGTAAATGAC | TGTAAGGCAAGAGGTTCC |
| Slco6c1 | AGTTCATGCGGCTTCTATTTA | GTTTTCCACCACCACTCTGC |
| Rsp16 | TCGGACGCAAGAAAACAGC | GACCCGAATATCCACACCAG |

Supplemental Table 4 – Real-time PCR primer list.