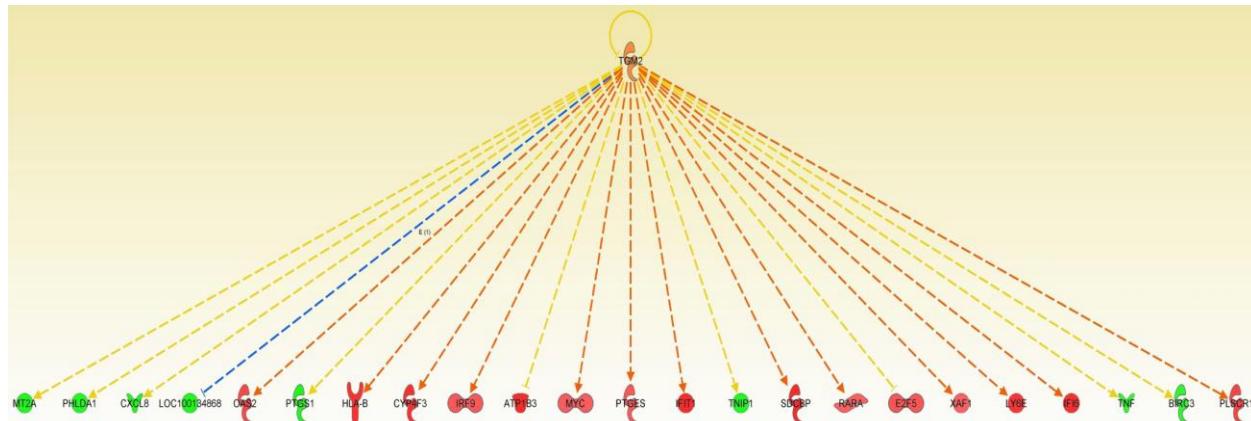


The expression of genes contributing to pancreatic adenocarcinoma progression is influenced by the respective environment – Sagini et al



Supplementary Figure 1: Target genes regulated by TGM2. Figure represents 24 genes regulated by TGM2, which were obtained from Ingenuity Pathway Analysis. As indicated, 9 genes (marked red) are down-regulated by TGM2. On the contrary, 15 genes (marked red) are up-regulated by TGM2.

Supplementary Table 1: Functional annotations of genes from Suit2-007 cells growing in pancreatic environment

| Categories ^a | Diseases or Functions Annotation ^b | p-Value ^c | Predicted activation ^d | Activation Z-score ^e | Number of genes ^f |
|--|---|----------------------|-----------------------------------|---------------------------------|---|
| Cell movement | Cell movement | 1,56E-11 | increased | 2,199 | LAMB3, CEACAM6, CCL20, AGR2, MUC1, CXCL1, LAMA3, LCN2, COL17A1, CXCL8, AIF1, MMP7, CEMIP, JUP, SOD2, S100A4, PDGFA, NDRG1, SGK1, IGFBP3, DDR1, IL1A, CDKN1A, NREP, SEMA3E SERPINA3, SDC4, ALPP, CX3CL1, NFKBIA, ANXA3, CDH1, CDCP1, CRYAB, TUBB2B, FOXQ1, SLPI, F3, GRINA, ITGA2, ARPIN/C15orf38-AP3S2, SPTLC1, IL10, TSC22D3, LAMC2, TCAF1, CDH3, MX1, LEP, ZC3H12A, PMP22, IL32, FAM83H, EFNA1, PATJ, CEBPB, SERPINA5, PTK6, EPHB6, JUND, TNFSF14, ERBB3, TNFRSF25, FCAR, CXCL16, HLA-A, CEACAM1, FAT1, AHR, CSF2RA, CLDN7, MAPK13, FERMT1, TCAF2, MST1R, CD99, PTP4A2, PHLDA1, DEFB1, RHOB, TNFSF15, CD44, CSF2, SERPINB5, TGM2, SRC, ITGA6, TNC, HNRNPA2B1, RHOD, SKI, KISS1, TACSTD2, GNAI2, CXCL2, NFKB2, TAGLN2, TNF, CD74, PTPRK, STAT3, ARHGAP21, VEGFA, MYH9, SAA1, F11R, PDCD4, IQGAP1, DCN, MAPK8IP3, STC1, ADAM15, LTBP2, HOOK1, CST3, EPHA1, TIMP2, LPAR2, CORO1A, CLDN3, MYO1C, ANGPTL4, SPHK2, NME1, COL18A1, PRMT1, FAF1, EZH2, SPRY2, LASP1, KITLG, CDK4, DPAGT1, LPAR1, F2R, NRP1, SELP, PBK, HNRNP, JUN, HMMR, TUBA1C, ST3GAL5, RUVBL2, ACP1, KLK3, CDKN3, CITED2, SKP2, IGF2BP3, KIAA1524, CAV1, NET1, CDX2, CDK1, ID2, CYR61, SDCBP, HSPD1, TGFB1, PPARG, ANXA2, TMPO, VIPR1, NR5A2, EGR1, RGS4, FOSB, DNAJB6, ADRB2, PTTG1, TPM3, MB, CTGF, KPNA2, TNFRSF1B, FOS, THBS (178/326) |
| Cell-to-Cell Signaling and Interaction | Interaction of tumor cell lines | 2,40E-10 | increased | 2,559 | MUC1, LAMA3, CXCL8, MUC16, IGFBP3, DDR1, CX3CL1, NFKBIA, CDH1, CDCP1, F3, ITGA2, LAMC2, CDH3, LEP, SERPINA5, EPHB6, ARID1A, ERBB3, TNFRSF25, HLA-A, CD99, PHLDA1, CD44, SERPINB5, TGM2, SRC, ITGA6, TNC, CD68, CXCL2, TNF, DYSF, STAT3, ARHGAP21, VEGFA, F11R, HYAL1, DCN, ADAM15, SPTAN1, NME1, PRMT1, KITLG, INHBB, F2R, SELP, JUN, HMMR, ACP1, KLK3, ERP29, CAV1, NET1, PPARG, ANXA2, VIPR1, MAD2L1, FOS, THBS(60/109). |
| Gene expression | Expression of RNA | 6,19E-09 | increased | 2,572 | AGR2, SOD2, PDGFA, IL1A, CDKN1A, IRF2BPL, BAMBI, KMT2E, APOBEC3G, KLK6, HOXA5, NFKBIA, FOXJ1, CDH1, CRCP, BLZF1, CREB1, NFIB, TRIM13, IL10, TSC22D3, FOXO4, AIRE, CDH3, ZC3H12A BHLHE40, CEBPB, ONECUT2, ARID1A, JUND, TNFSF14, ZNF148, TNFRSF25, IRF9, JAG1, CAPRIN2, AHR, MAGED1, TSC22D1, MNT, MYBL1, BTG1, LRRKIP1, TOB1, STAT2, FARS2, HNF1B, RHOB, GABPB2, TNFSF15, CD44, LILRB1, CSF2, TGM2, SRC, TNIP1, ITGA6, HNRNPA2B1, RBM14, DNNTIP1, SKI, CFLAR, DDIT3, FLII, LDOC1, NFAT5, NFKB2, TNF, PTPRK, STAT3, NFIX, EIF4A2, GSN, VEGFA, MAFF, PDCD4, IQGAP1, ADIRF, DCN, TNFAIP3, MED15, KLF9, LPAR2, UBR5, CBX2, TAF9, NME1, FAF1, EZH2, SPRY2, MELK, ATAD2, NKX3-1, ZFP36, INHBB, LPAR1, F2R, SFR1, HNRNP, JUN, RUVBL2, ARF4, PHGDH, RBM4, RXRA, UHRF1, TIPARP, PLK1, |

| | | | | | |
|---|--|----------|-----------|--------|--|
| | | | | | ERP29, SUV39H1, CITED2, ARID5B, MED27, PAGR1, IGF2BP3, GMNN, BIRC5, CAV1, GTF2A2, PSRC1, TXNRD1, EIF5, CDK2, TRIP13, CDX2, ID2, CYR61, RAE1, PPARG, DHX9, NR5A2, STC2, CCND3, EGR1, FOXL2, FZD2, FOSB, PTTG1, RPL6, HNRNPA1, ALYREF, TNFRSF1B, FOS (143/341) |
| Cellular assembly & organization, DNA replication, recombination & repair | Formation of nuclear foci | 3,33E-05 | increased | 3,142 | CCNB1, NKX3-1, RUVBL2, RFWD3, CDK1, CCND3(6/24) |
| Cell death & survival | Cell death of squamous cell carcinoma cell lines | 4,18E-05 | increased | 2,414 | JUP, CD44, KISS1, CFLAR, DDIT3, TNF, STAT3, PDCD4, PLK1, BIRC5, RRM2, FBXO5 (12/24) |
| Cell death & Survival | Cell viability of ovarian cancer cell lines | 1,69E-04 | increased | 2,215 | SOD2, SLPI, JAG1, HNF1B, SRC, CLDN3, JUN (7/17) |
| Cancer, organismal injury & abnormalities | Growth of tumor | 2,18E-07 | decreased | -2,198 | MUC1, IGFBP3, CDKN1A, BAMBI, ZMAT3, IL10, LEP, BIK, CEACAM1, JAG1, CSF2RA, SSTR2, MST1R, RHOB, CD44, CSF2, SERPINB5, SRC, SKI, DDIT3, TNF, STAT3, VEGFA, PTPRA, DCN, ADAM15, LPAR2, PPP2CA, KITLG, NKX3-1, INHBB, CDK4, KLK3, RXRA, PLK1, PEG10, SKP2, BIRC5, CAV1, ID2, CYR61, EGR1, SLC3A2, FOS, THBS1 (45/59) |
| Infectious diseases | Viral infection | 1,44E-12 | decreased | -2,198 | HBA1/HBA2, LCN2, CXCL8, CLIP3, S100A4, SLC2A3, SGK1, ANO1, IL1A, IL23A, ALPP, APOBEC3G, NFKBIA, BIRC3, HMGCS1, MAGT1, TUBA3C/TUBA3D, SLPI, HIST1H2BD, CREB1, HNRNPU, CRYAB, F3, PDE4C, ARL16, IL10, MX1, TMEM132A, LEP, ZC3H12A, IL32, ARHGAP32, NUBPL, SERPINA5, PLOD2, EPHB6, ARID1A, HIST2H2AA3/HIST2H2AA4, ZNF148, IRF9, HLA-A, CSF2RA, AP2B1, MST1R, TNFRSF14, WNT10A, CLK1, CYP3A5, PSMD12, STAT2, DEFB1, RHOB, GABPB2, F5, SHCBP1, AP1G2, CD44, CSF2, BAG6, OPTN, KANSL1, PDE4A, HSPA6, FUT2, TCIRG1, HLA-B, FLII, TAGLN2, TNF, PSENEN, CD74, DYSF, STAT3, ARHGAP21, F11R, SRRM2, DUSP3, PPM1K, PTGS1, LDLR, COPG1, MED15, STXBP2, CST3, HPGD, SPTAN1, TIMP2, TMEM63A, LPAR2, CLDN3, NUDCD3, UBR5, DCLK1, PDXK, POLR1E, SNRPF, CDC42EP3, ERCC1, ADK, PDSS1, MND1, HSPA4, EZH2, SPRY2, SF3B6, SRRT, VNN2, SAFB, CDK4, F2RNDFUF2, PBK, DIMT1, TKT NUDT11, HNRNPK, JUN, TUBA1C, ST3GAL5, RUVBL2, G3BP2, EXOSC3, NCKAP1, MT2A, RXRA, PLK1, PSMA3, TUBG1, UBE2L6, MED27, SNU13, CAV1, PSMA1, UNG, CDK2, RRM2, KRT18, HSPD1, PPARG, RANBP1, DHX9, MGLL, ANXA2, ARHGAP23, TYMS, CAV2, AP1S1, TUBB, TUBA1A, EGR1, UBE2E1, DUSP1, ADRB2, PTTG1, GINS4, PSMC4, TNFRSF1B (156/362) |
| Cellular | Alignment of | 6,68E-06 | decreased | -2,976 | NCAPG2, SMC4, KIFC1, KIF20A, BIRC5, KIF2C, DLGAP5, SGO1 (8/12) |

| | | | | | |
|---|--|----------|-----------|--------|---|
| assembly & organization, DNA replication, recombination, and repair | chromosomes | | | | |
| Cell cycle | M phase of tumor cell lines | 6,89E-05 | decreased | -2,240 | SVIL, GNAI2, STX16, CDT1, PLK1, BIRC5, KIF20A, KIFC1, TRIP13, FBXO5, MAD2L1, PTTG1, MCM7, SPDL1(14/29) |
| Cell cycle | Cell cycle progression of cervical cancer cells | 6,93E-05 | decreased | -2,138 | HYPK, AHR, PLK1, BIRC5, BANF1, NET1, CDK1, MAD2L1, TCP1 (9/29) |
| DNA replication, recombination, and repair | Double-stranded DNA break repair | 9,29E-05 | decreased | -2,135 | FAM175A, TDP1, ARID1A, TNKS1BP1, ERCC1, EZH2, G6PD, EXO1, SUV39H1, CDK1, KPNA2(11/30) |
| Cell cycle | Interphase of bone cancer cells | 9,50E-05 | decreased | -2,343 | NDRG1, CDH1, LEP, JAG1, MST1R, CD44, TGM2, TNF, STAT3, ARHGAP21, VEGFA, HOOK1, PRMT1, HNRNPAB, NRP1, CAV1, PTTG1, CTGF(18/20) |
| Cellular development | Epithelial-mesenchymal transition of tumor cells | 1,05E-04 | decreased | -2,678 | CDKN1A, BHLHE40, NEK11, ERCC1, PLK1, CDK2 (6/10) |
| Cell cycle | Re-entry into interphase | 1,09E-04 | decreased | -2,121 | KIFC1, NUSAP1, KIF20A, BIRC5, PLK1, NME1, CCNB1, STX16, MYH9, GNAI2, SVIL, RHOB, INF2, CDKN1A (15/36) |

^a Categories, are broader functions under which various gene cluster a were annotated by Ingenuity pathway analysis.

^b Diseases or functions annotations represent predicted biological functions of gene clusters derived Ingenuity Pathway Analysis.

^cp-Value is the calculated probability derived from individual gene cluster in a defined annotation for the respective gene group.

^dPredicted activation is the overall activation state (increased or decreased) of the functional annotation for respective genes in a given gene cluster.

^eActivation Z-score is a defined quantity, which determines whether a biological function has significantly more “increased” predictions than “decreased” predictions ($Z>0$) or vice versa ($Z<0$).

^fNumber of genes represent selected genes based on significant modulation of expression versus the total number of genes for respective functional annotation.

Supplementary Table 2: Functional annotations of genes from Suit2-007 cells growing in liver environment

| Categories ^a | Diseases or Functions Annotation ^b | p-Value ^c | Predicted activation ^c | Activation Z-score ^e | Number of genes ^f |
|---|---|----------------------|-----------------------------------|---------------------------------|--|
| Cell-to-cell signaling & interaction | Interaction of tumor cells | 1,23E-13 | increased | 2,191 | LGALS4, KLK5, RHOU, DCN, LGALS2, APP, NR1H4, KITLG, CXCR4, CAV1, ERBB3, TNFRSF25, ITGB1, ANXA2, CDH1, HYAL1, ITGB2, ALCAM, SDC1, ARID1A, LAMA3, PKM, PHLDA1, TNFRSF6B, IRS1CDCP1, EPHB6, CLDN2, LAMC2 (28/77) |
| Cellular movement, immune cell trafficking | Leukocyte migration | 3,58E-05 | increased | 3,039 | DEFB1, GDF15, CD74, S100A9, PPBP, CRP, SERPINA1, APP, KITLG, CXCR4, CXCL6, EDNRB, AIF1, CCL20, SAA1, ITGB1, ITGB2, ALCAM, ENG, ADORA1, LGALS1, TNFRSF6B, IL1A(23/49) |
| Inflammatory response | Inflammatory response | 5,19E-06 | increased | 2,423 | DEFB1, CD74, S100A9, PPBP, CRP, SERPINA1, APP, KITLG, CXCR4, CXCL6, SERPING1, EDNRB, AIF1, CCL20, SAA1, HYAL1, MGLL, ITGB2, TNFAIP6, ENG, ADORA1, LGALS1, TNFRSF6B, IL1A, ADAM8(25/49) |
| Cell-to-cell signaling and interaction | Activation of cells | 3,91E-05 | increased | 2,011 | LGALS4, HLA-DPA1, CRP, APP, B2M, CXCR4, CXCL6, SERPING1, ATP2A2, CDH1, TNFSF10, ITGB2, PKM, TNFRSF6B, IL23A, IL1A, LTB (17 /53) |
| Cellular movement | Homing of cells | 1,84E-04 | increased | 2,998 | DEFB1, CD74, RHOU, S100A9, PPBP, SERPINA1, APP, KITLG, CXCR4, CXCL6, MDK, EDNRB, AIF1, CCL20, CAV1, SAA1, ITGB1, ANXA2, ITGB2, ADORA1, LGALS1, TNFRSF6B, FOSL1 (23/48) |
| Cellular growth and proliferation | Colony formation of breast cancer cell lines | 5,90E-04 | increased | 2,044 | FGFR3, SERPINA1, ELF3, IL1A (4/15) |
| Neurological, skeletal & muscular disorders | Neuromuscular disease | 5,54E-05 | increased | 2,449 | FGG, HLA-DRA, GABRP, HLA-DRB1, CD74, FGFR3, ALDH6A1, MT1H, HLA-DQA1, HLA-B, CRP, SERPINA1, IFIT1, APP, USP2, HLA-DMA, SERPING1, ALOX5, ETFB, ATP2A2, DPAGT1, GLRB, MX1, PLSCR4, FOXN3, DNAJB6, ISG15, ANXA2, NDRG2, HNRNPDL, TUBB2B, ALCAM, LY6E, ADRA1B, PKM, CSRP2, LAMB1, ADORA1, ARRB1, MAOB, AP1S2, PTGS1, IL1A, COL13A1 (48/106) |
| Neurological disease | Progressive motor neuropathy | 3,02E-07 | increased | 2,449 | HLA-DRA, TSPAN8, NNMT, GABRP, HLA-DRB1, CD74, FGFR3, HLA-DQA1, HLA-B, IFIT1, APP, PDGFC, FUCA1, HLA-DMA, SERPING1, ALOX5, MX1, DNAJB6, ISG15, ANXA2, GADD45A, NDRG2, HNRNPDL, SHROOM3, TUBB2B, ALCAM, LY6E, ADRA1B, ADORA1, ADIRF, ARRB1, MAOB, PFKP, AP1S2, PTGS1, IL1A (36/81) |

| | | | | | |
|-------------------------|------------|----------|-----------|--------|---|
| Cell death and survival | Cell death | 3,68E-16 | decreased | -2,162 | GSTA2, LGALS4, GSTA1, DEFB1, IFI27, GABRP, HLA-DRB1, GDF15, CD74, FGFR3, S100A9, LMO4, PPBP, HLA-DRB4, LCN2, SEMA6A, LGALS2, CRP, APP, NR1H4, USP2, KITLG, B2M, PI3, IFI6, PDGFC, CXCR4, MMP7, IGFBP7, MDK, ALOX5, MT1F, ARF4, ATP2A2, ITM2B, ID2, EDNRB, NAMPT, SCIN, IFIT2, NDRG1, PPM1A, HBA1/HBA2, CHMP5, WSB1, PLSCR1, MX1, TCP1, KLK6, CAV1, ANTXR1, AKR1C3, NFKBIZ, SAA1, ERBB3, TNFRSF25, ITGB1, ISG15, ADGRL2, ANXA2, ANXA4, CDH1, GADD45A, WDR19, HYAL1, NDRG2, TNFSF10, ADM, PDZK1, ITGB2, NEK11, MYC, ALCAM, GPR37, ZFAND6, DUSP1, SDC1, KRT18, ENG, THOC2, HIPK2, PKM, PHLDA1, ADORA1, LGALS1, TNFRSF6B, SULF2, PHLDA2, CAPRIN2, LMNA, ARRB1, IRS1, FOSL1, MAOB, MLLT11, TRPV2, CDC25B, CDCP1, PMP22, EPHB6, LOC102724788/PRODH, PTGS1, IL1A (103/306) |
|-------------------------|------------|----------|-----------|--------|---|

^aCategories are broader functions under which various gene cluster are annotated by Ingenuity pathway analysis;

^bCategories or Functions annotations represent predicted biological functions of gene clusters derived Ingenuity Pathway Analysis

^cp-Value is the calculated probability derived from individual gene cluster in a defined annotation for the respective gene group.

^dPredicted activation is the overall activation state (increased or decreased) of the functional annotation for respective genes in a given gene cluster

^eActivation Z-score ; is a defined quantity, which determines whether a biological function has significantly more “increased” predictions than “decreased” predictions ($Z>0$) or vice versa ($Z<0$).

^fNumber of genes, represent selected genes based on significant modulation of expression versus the total number of genes for respective functional annotation

Supplementary Table 3: Functional annotations of genes from Suit2-007 cells growing in lung environment

| Categories ^a | Diseases or Functions Annotation ^b | p-Value | Predicted activation | Activation Z-score ^c | Number of genes ^d |
|---|---|----------|----------------------|---------------------------------|---|
| Cell death & survival | Apoptosis | 1,80E-16 | decreased | -2,144 | LGALS4, IFI27, GSTA2, EDNRB, HOXA5, GSTA1, FGFR3, CRYAB, LGALS2, ID2, PPP1R1B, APP, LMO4, BDNF, DEFB, GDF15, AGT, EIF4G2, PRDX3, ARF4, TCP1, ITGB1, ASNS, EHF, SON, MX1, ATP1B3, CD24, ZFAND6, FOXO4, B2M, CTNNB1, UBA3, DPP4, NET1, PPM1A, BEX2, AQP1, IL1AANXA4, DKK1, DPEP1, IFI6, JUND, CDH1, FHL2, RPS6KA3, MSN, LGALS1, CSF2, KISS1, CLIP3, IGFBP6, TGM2, CXCL1, ANGPTL4, IGFBP3, EPHB6, CXCL8, CDCP1(59/175) |
| Cell death & survival | Cell viability of tumor cell lines | 6,69E-07 | increased | 2,657 | HBA1/HBA2, HOXA5, FGFR3, ID2, PPP1R1B, APP, BDNF, ARID1A, GDF15, TCP1, ITGB1, ASNS, CD24, CTNNB1, DPP4, PSMC4, BEX2, CDH1, RPS6KA3, DNAJB6, ARRB1, MDK, HSPA6, NFAT5, CSF2, TGM2, IGFBP3, CXCL8, IL1A (29/77) |
| Cellular growth, development, & proliferation | Cell proliferation of tumor cell lines | 1,06E-13 | increased | 2,979 | ADAM10, APP, CDH17, CXCL1, ERBB3, GABRP, GSK3B, IGFBP3, ILK, LGALS1, LOX, MYC, NAMPT, OGT, RRM2, STAT3, TMPO, TNF, WSB1(19/157) |
| Organismal development | Growth of organism | 1,80E-04 | increased | 2,461 | AMD1, APP, BMP4, CDC42, CNOT2, CRCP, CTNNB1, DEFB1, DICER1, DKK1, GSK3B, GSN, H19, IGFBP3, MID1IP1, PNPT1, SLC30A5, SUMO2, TNF (19/19) |
| Gene expression | Transactivation | 8,91E-07 | increased | 2,607 | APP, LMO4, BDNF, ARID1A, AGT, EHF, MX1, CTNNB1, DPP4, BEX2, ANXA4, THRA, CDH1, FHL2, NFIB, HERC5, ARRB1, LGALS1, CXCL8, IL1A (20/46) |
| Lipid metabolism & molecular transport | Release of fatty acid | 1,54E-03 | decreased | -2,176 | BDNF, CSF2, IL1A, IL32, KISS1, LEP, LGALS1, PTGS1, TNF (9/9) |
| Infectious diseases | Viral infection | 1,40E-04 | increased | 2,953 | HBA1/HBA2, CRYAB, DEFB1, ARID1A, IFIT1, PSENEN, TCN1, SPTBN1, ITGB1, CAV2, SON, MX1, CD24, PICALM, B2M, CTNNB1, DPP4, PSMC4, NUP153, ANXA2, EFNB2, SUMO2, DKK1, HCP5, HLA-B, RPS6KA3, LY6E, ARRB1, HSPA6, SLC2A3, LGALS1, CSF2, HNRNPU, ARL16, MAGT1, CLIP3, APOBEC3G, EPHB6, CXCL8, IL1A, IL23A (41/116) |
| Cellular movement | Invasion of melanoma cell lines | 1,03E-03 | decreased | -2,157 | BMP4, CDH1, CXCL8, FXYD5, ID2, MMP1, MX1, NFAT5, SDCBP, STAT3 (10/10) |

^a Categories are broader functions under which various gene cluster are annotated by Ingenuity pathway analysis.

^bDiseases or functions annotations represent predicted biological functions of gene clusters derived Ingenuity Pathway Analysis.

^cp-Value is the calculated probability derived from individual gene cluster in a defined annotation for the respective gene group.

^dPredicted activation is the overall activation state (increased or decreased) of the functional annotation for respective genes in a given gene cluster.

^eActivation Z-score is a defined quantity, which determines whether a biological function has significantly more “increased” predictions than “decreased” predictions ($Z>0$) or vice versa ($Z<0$).

^fNumber of genes represent selected genes based on significant modulation of expression versus the total number of genes for respective functional annotation.

Supplementary Table 4: RT-PCR experiment following siRNA knockdown of TGM2

| TGM2-24 h | Cp 1 | Cp 2 | Cp 3 | Average | Average GAPDH | Normalized | Average control | Cp difference. | Expression | % Kd. |
|------------------------|-------|-------|-------|---------|---------------|------------|-----------------|----------------|------------|-------|
| Control | 29.26 | 30.26 | 28.93 | 29.48 | 14.32 | 15.16 | 15.11 | | | |
| Control + Lipid | 29.57 | 29.52 | 28.8 | 29.30 | 14.23 | 15.06 | | | | |
| 1:4 , 50nM | | 31.57 | | 31.57 | 14.72 | 16.85 | | 1.74 | 0.30 | 70 |
| 1:5 , 50nM | 29.91 | 30.4 | 29.57 | 29.96 | 14.37 | 15.59 | | 0.47 | 0.72 | 28 |
| 1:4 , 100nM | 34.19 | 33.73 | 34.66 | 34.19 | 14.79 | 19.41 | | 4.29 | 0.05 | 95 |
| TGM2-48 h | Cp 1 | Cp 2 | Cp 3 | Average | Average GAPDH | Normalized | Average control | Cp difference | Expression | % Kd. |
| Control | 30.84 | 29.9 | 30.33 | 30.36 | 14.79 | 15.57 | 15.42 | | | |
| Control + Lipid | 30.96 | 29.79 | 29.5 | 30.08 | 14.81 | 15.28 | | | | |
| 1:4 , 50nM | | 30.75 | 31.01 | 30.88 | 14.58 | 16.30 | | 0.88 | 0.54 | 46 |
| 1:5 , 50nM | 32.51 | 31.72 | 33.36 | 32.53 | 14.93 | 17.60 | | 2.18 | 0.22 | 78 |
| 1:4 , 100nM | 34.07 | | 33.13 | 33.60 | 14.85 | 18.75 | | 3.33 | 0.10 | 90 |

Table4 represent results of RT-PCR experiment following TGM2 knockdown in suit2-007 cells with different siRNA concentrations at 24 h and 48 h, respectively. Shown are the average Cp (crossing point) values for each triplicate treatment. The values for GAPDH (control) are also shown. The percent knockdown (% kd) of TGM2is shown in the last column.

Supplementary table5:

Selected genes and their respective proteins (Chip array experiment)

| Gene Name | Protein name |
|-----------|---|
| ACP1 | Adipocyte acid phosphatase) phosphatase 1 |
| ACTN1 | Alpha-actinin-1 |
| ADAM8 | Disintegrin and metalloproteinase domain-containing protein 8 (ADAM 8) |
| ADD3 | Gamma-adducin (Adducin-like protein 70) |
| ADORA1 | Adenosine receptor A1 |
| AGR2 | Anterior gradient protein 2 homolog (AG-2) |
| AGT | Angiotensinogen (Serpine A8) |
| AIF1 | Allograft inflammatory factor 1 (AIF-1) |
| ANXA1 | Annexin A1 |
| ANXA2 | Annexin A2 (Annexin II) (Annexin-2) |
| ANXA4 | Annexin A4 (35-beta calcimedin) (Annexin IV) |
| APOBEC3G | DNA dC->dU-editing enzyme APOBEC-3G |
| APP | Amyloid-beta A4 protein (ABPP, APP1, APP) |
| AQP1 | Aquaporin-1 (AQP-1) |
| AQP5 | Aquaporin-5 (AQP-5) |
| ARF4 | ADP-ribosylation factor 4 |
| ARHGEF1 | Rho guanine nucleotide exchange factor 1 (p115RhoGEF) |
| ARID1A | AT-rich interactive domain-containing protein 1A |
| ARL6IP1 | ARL-6-interacting protein 1 (Aip-1), Apoptotic regulator in the membrane of the endoplasmic reticulum |

| | |
|---------|--|
| ARPC3 | Actin-related protein 2/3 complex subunit 3 |
| ARRB1 | Beta-arrestin-1 (Arrestin beta-1) |
| ATM | Serine-protein kinase ATM |
| ATP1B3 | Sodium/potassium-transporting ATPase subunit beta-3 |
| B2M | Beta-2-microglobulin |
| BCL2L1 | Bcl-2-like protein 1 (Bcl2-L-1), Apoptosis regulator Bcl-X |
| BDNF | Brain-derived neurotrophic factor (BDNF), Abrineurin |
| BHLHE40 | Class E basic helix-loop-helix protein 40 (bHLHe40) |
| BIRC2 | Baculoviral IAP repeat-containing protein 2 |
| BRE | BRISC and BRCA1-A complex member 2 |
| CAPRIN2 | Caprin-2 (C1q domain-containing protein 1) |
| CAV1 | Caveolin1 |
| CCL20 | C-C motif chemokine 20 (Beta-chemokine exodus-1) |
| CCND3 | G1/S-specific cyclin-D3 |
| CD151 | CD151 antigen (GP27), Membrane glycoprotein SFA-1 |
| CD58 | CD antigen CD58) |
| CD74 | HLA class II histocompatibility antigen gamma chain (HLA-DR), CD antigen CD74) |
| CDC20 | Cell division cycle protein 20 homolog (p55CDC) |
| CDC42 | Cell division control protein 42 homolog |
| CDCP1 | CUB domain-containing protein 1 (Membrane glycoprotein gp140) |
| CDH1 | Cadherin-1 (CAM 120/80), Epithelial cadherin, E-cadherin |
| CDK2 | Cyclin-dependent kinase 2 |
| CDKN1A | Cyclin-dependent kinase inhibitor 1 |
| CHMP5 | Charged multivesicular body protein 5 (Chromatin-modifying protein 5) |
| CLDN2 | Claudin-2 (SP82) |
| COL13A1 | Collagen alpha-1(XIII) chain (COLXIIIA1) |
| CRYAB | Alpha-crystallin B chain (Alpha(B)-crystallin) |
| CSF2 | Colony-stimulating factor) (CSF) |
| CSF2RA | Granulocyte-macrophage colony-stimulating factor receptor subunit alpha |
| CTGF | Connective tissue growth factor (CCN family member 2) |
| CTNNB1 | Catenin beta-1 (Beta-catenin) |
| CXCL16 | C-X-C motif chemokine 16 |

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| CXCL8 | Interleukin-8 (IL-8) |
| DCN | Decorin (Bone proteoglycan II) |
| DDIT3 | DNA damage-inducible transcript 3 protein (DDIT-3) |
| DEFB1 | Beta-defensin 1 (BD-1), Defensin beta 1 |
| DPP4 | Dipeptidyl peptidase 4 (ADABP) |
| DYSF | Dysferlin (Dystrophy-associated fer-1-like protein) |
| EDNRB | Endothelin receptor type B (ET-B) |
| EHF | ETS homologous factor (hEHF) |
| EIF4G2 | Eukaryotic translation initiation factor 4 gamma 2 |
| ENG | Endoglin (CD antigen CD105) |
| EPCAM | Epithelial cell adhesion molecule (Ep-CAM) |
| EPHB6 | Ephrin type-B receptor 6 (HEP) |
| ERBB3 | Receptor tyrosine-protein kinase erbB-3 |
| ERP29 | Endoplasmic reticulum resident protein 29 (ERp29) |
| ETS1 | Protein C-ets-1 (p54) |
| FBXO5 | F-box only protein 5 (Early mitotic inhibitor 1) |
| FGFR3 | Fibroblast growth factor receptor 3 (FGFR-3) |
| FGG | Fibrinogen gamma chain |
| FHL2 | Four and a half LIM domains protein 2 (FHL-2) |
| FOS | Proto-oncogene c-Fos (Cellular oncogene fos) |
| FOSL1 | Fos-related antigen 1 (FRA-1) |
| GADD45A | Growth arrest and DNA damage-inducible protein GADD45 alpha |
| GDF15 | Growth/differentiation factor 15 (GDF-15) |
| GSK3B | Glycogen synthase kinase-3 beta (GSK-3 beta) |
| GSTA1 | Glutathione S-transferase A1 |
| GSTA2 | Glutathione S-transferase A2 |
| HAT1 | Histone acetyltransferase 1 |
| HERPUD1 | Homocysteine-responsive endoplasmic reticulum-resident ubiquitin-like domain member 1 protein |
| HLA-A | HLA class I histocompatibility antigen, |
| HLA-DPA1 | HLA class II histocompatibility antigen, DP alpha 1 chain |
| HLA-DRA | HLA class II histocompatibility antigen, DR alpha chain |
| HNRNPA1 | Heterogeneous nuclear ribonucleoprotein A1 (hnRNP A1) |
| HNRPK | Heterogeneous nuclear ribonucleoprotein K (hnRNP K) |
| HOXA5 | Homeobox protein Hox-A5 |

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| HYAL1 | Hyaluronidase-1 (Hyal-1) |
| HYPK | Huntingtin-interacting protein K (Huntingtin yeast partner K) |
| ID2 | DNA-binding protein inhibitor ID-2 |
| IER3 | Radiation-inducible immediate-early gene IEX-1 |
| IFI27 | Interferon alpha-inducible protein 27 |
| IFI6 | Interferon alpha-inducible protein 6 |
| IFIT1 | Interferon-induced protein with tetratricopeptide repeats 1 (IFIT-1) |
| IGF2 | Insulin-like growth factor II (IGF-II) |
| IGFBP3 | Insulin-like growth factor-binding protein 3 (IBP-3) |
| IL1A | Interleukin-1 alpha (IL-1 alpha) |
| IL23A | Interleukin-23 subunit alpha (IL-23 subunit alpha) |
| IL32 | Interleukin-32 (IL-32) (Natural killer cells protein 4) |
| IL8 | Interleukin-8 (IL-8) (C-X-C motif chemokine 8) |
| INF2 | Inverted formin-2 (HBEBP2-binding protein C) |
| IRS1 | Insulin receptor substrate 1 (IRS-1) |
| ITGB1 | Integrin beta-1 (Fibronectin receptor subunit beta) |
| ITGB2 | Integrin beta-2 |
| JAK1 | Tyrosine-protein kinase JAK1 |
| JUN | Proto-oncogene c-Jun |
| JUND | Transcription factor jun-D |
| JUP | Junction plakoglobin (Catenin gamma) |
| KCNN4 | Intermediate conductance calcium-activated potassium channel protein 4 (SK4) |
| KISS1 | Metastasis-suppressor KiSS-1 (Kisspeptin-1) |
| KITLG | Kit ligand (Mast cell growth factor) |
| KLF9 | Krueppel-like factor 9 |
| KPNA2 | Importin subunit alpha-1 (Karyopherin subunit alpha-2) |
| LAMA3 | Laminin subunit alpha-3 (Epiligrin 170 kDa subunit) |
| LAMB3 | Laminin subunit beta-3 (Epiligrin subunit bata) |
| LAMC2 | Laminin subunit gamma-2 (Cell-scattering factor 140 kDa subunit) |
| LGALS1 | Galectin-1 (Gal-1), (Beta-galactoside-binding lectin L-14-I) |
| LGALS2 | Galectin-2 (Gal-2), Beta-galactoside-binding lectin L-14-II |
| LGALS4 | Galectin-4 (Gal-4), Lactose-binding lectin 4 |
| LMO4 | LIM domain transcription factor LMO4 |
| LTB | Lymphotoxin-beta (LT-beta) |

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| LTBR | Lymphotoxin-beta receptor, Tumor necrosis factor C receptor |
| MAOA | Monoamine oxidase type A (MAO-A) |
| MAX | Protein max (Class D basic helix-loop-helix protein 4) |
| MDK | Midkine (MK), Amphiregulin-associated protein |
| MGAT3 | Beta-1,4-mannosyl-glycoprotein 4-beta-N-acetylglucosaminyltransferase |
| MMP1 | Matrix metalloproteinase-1 |
| MSN | Moesin (Membrane-organizing extension spike protein) |
| MUC1 | Mucin-1 (MUC-1) |
| MUC16 | Mucin-16 (MUC-16) |
| MX1 | Myxovirus resistance protein 1 |
| MYC | Myc proto-oncogene protein |
| NAMPT | Nicotinamide phosphoribosyltransferase (NAmPRTase) |
| NCAPG2 | Condensin-2 complex subunit G2 |
| NCKAP1 | Nck-associated protein 1 (NAP 1) |
| NDRG1 | Protein NDRG1 (Differentiation-related gene 1 protein) |
| NFAT5 | Nuclear factor of activated T-cells 5 (NF-AT5) |
| NFIB | Nuclear factor 1 B-type (NF1-B) |
| NR1H4 | Bile acid receptor , (Nuclear receptor subfamily 1 group H member 4) |
| PDGFC | Platelet-derived growth factor C (PDGF-C) |
| PHLDA1 | Pleckstrin homology-like domain family A member 1 |
| PKM2 | Pyruvate kinase PKM |
| PLAT | Tissue-type plasminogen activator (t-PA) chain B] |
| PLSCR1 | Phospholipid scramblase 1 (PL scramblase 1) |
| PPARG | Peroxisome proliferator-activated receptor gamma |
| PPM1A | Protein phosphatase 1A (EC 3.1.3.16) |
| PRDX3 | Thioredoxin-dependent peroxide reductase, mitochondrial |
| PRKCDBP | Protein kinase C delta-binding protein |
| PRNP | Alternative prion protein (AltPrP) |
| PRNP | Major prion protein (PrP) (ASCR) |
| PRODH | Proline dehydrogenase 1, mitochondrial |
| PTGS1 | Prostaglandin G/H synthase 1 |
| PTPN13 | Tyrosine-protein phosphatase non-receptor type 13 |
| PXN | Paxillin |
| RBBP4 | Histone-binding protein RBBP4 |

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| RHOU | Rho-related GTP-binding protein RhoU |
| RPS6KA3 | Ribosomal protein S6 kinase alpha-3 |
| RRM2 | Ribonucleoside-diphosphate reductase subunit M2 |
| SAA1 | Serum amyloid A-1 protein (SAA) |
| SDCBP | Syntenin-1 (Melanoma differentiation-associated protein 9) |
| SELP | P-selectin (CD62 antigen-like family member P) |
| SELP | Selenoprotein P (SeP) |
| SERPINA5 | Plasma serine protease inhibitor (Serpин A5) |
| SERPINB5 | Serpin B5 (Maspin) |
| SLC3A2 | Solute carrier family 3 member 2 |
| SLPI | Antileukoproteinase (ALP) (BLPI) |
| SOD2 | Superoxide dismutase [Mn], mitochondrial |
| SON | Protein SON (Bax antagonist selected in saccharomyces 1) |
| SPDL1 | Protein Spindly (hSpindly) |
| SPTLC1 | Serine palmitoyltransferase 1 |
| SSTR2 | Somatostatin receptor type 2 (SS-2-R) |
| STAT3 | Signal transducer and activator of transcription 3 |
| SULF2 | Extracellular sulfatase Sulf-2 (hSulf-2) |
| SVIL | Supervillin (Archvillin) |
| TCF2 | Transcription factor 2, (TCF-2) |
| TCP1 | T-complex protein 1 subunit alpha (TCP-1-alpha) |
| TGM2 | Transglutaminase-2, (TGase-2) |
| THBS1 | Thrombospondin-1 |
| TMED10 | Transmembrane emp24 domain-containing protein 10 |
| TNF | Tumor necrosis factor (Cachectin), TNF-alpha |
| TNFAIP3 | Tumor necrosis factor alpha-induced protein 3 |
| TNFRSF1B | Tumor necrosis factor receptor superfamily member 1B |
| TNFRSF25 | Tumor necrosis factor receptor superfamily member 25 |
| TNFRSF6B | Tumor necrosis factor receptor superfamily member 6B |
| TNFSF10 | Tumor necrosis factor ligand superfamily member 10 |
| TPT1 | Translationally-controlled tumor protein (TCTP) |
| TRIB3 | Tribbles homolog 3 (TRB-3) |
| TRPV2 | Transient receptor potential cation channel subfamily V member 2 |
| UBA3 | NEDD8-activating enzyme E1 catalytic subunit |

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| VIPR1 | Vasoactive intestinal polypeptide receptor 1 (VIP-R-1) |
| WSB1 | WD repeat and SOCS box-containing protein 1 |
| XAF1 | XIAP-associated factor 1 (BIRC4-binding protein) |
| ZFAND6 | AN1-type zinc finger protein 6 |
| ZFP36 | Zinc finger protein 823 (Zinc finger protein ZFP-36) |
| ZFP36 | mRNA decay activator protein ZFP36 |

Table 5 contains the names of genes selected from chip array experiment. These genes were evaluated based on their functional annotations as discussed in result section.