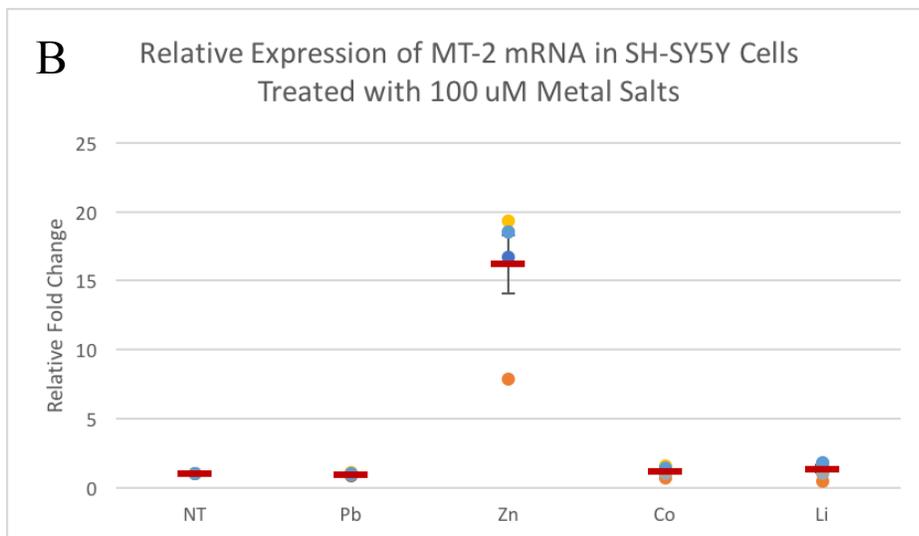
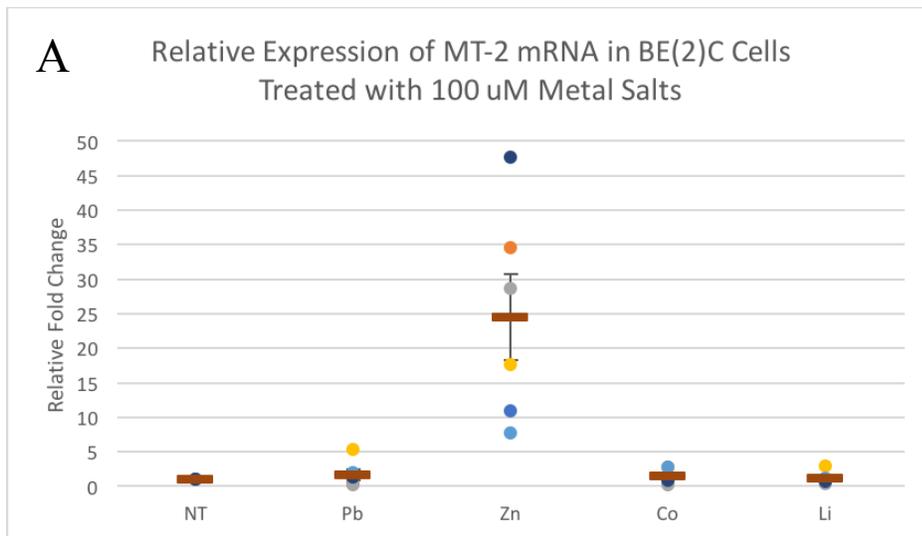
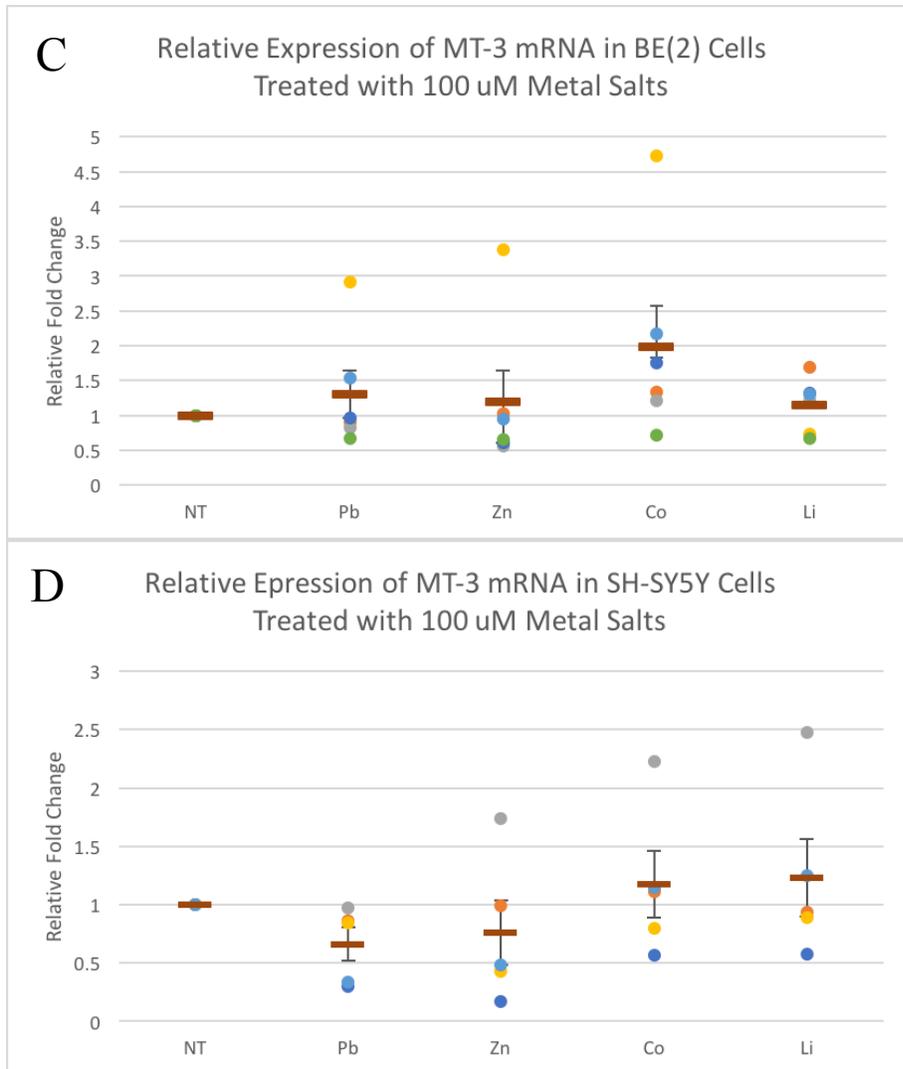


## Supplementary Materials: Function of Metallothionein-3 in Neuronal Cells: Do Metal Ions Alter Expression Levels of MT3?

Jamie Bousleiman, Alexa Pinsky, Sohee Ki, Angela Su, Irina Morozova, Sergey Kalachikov, Amen Wiqas, Rae Silver, Mary Sever, and Rachel Narehood Austin





**Figure S1. Relative fold change in cDNA expression after various metal treatments.** NT = no metal treatment. (A) *MT2* from BE(2)C cells; (B) *MT2* from SH-SY5Y cells; (C) *MT3* from BE(2)C cells; (D) *MT2* from SH-SY5Y cells. Average fold change is shown as a horizontal brown line, and standard deviation is shown as a vertical gray line.

**Table S1. Genes for metal-binding and Zn-binding GO categories found among upregulated genes in Group A and Group B.** Enriched terms for *Metal binding*, *Zinc-finger proteins* and *Zinc* were the same for both groups. **Count** is the number of differentially expressed genes matching the term; **%** is the percentage of the genes matching the term with respect to all differentially expressed genes that can be functionally annotated for the given group; **Pval** is a modified *Fisher Exact* test *P*-value for gene-enrichment analysis estimated using EASE.<sup>1</sup> *P*-values smaller than 0.05 indicate a strong enrichment in the functional annotation term.

Group A				Group B				
Term	Count	%	PVal	Genes	Count	%	PVal	Genes
<b>Metal-binding</b>	248	17	1.24E-05	PNMA3, DZIP1, OVCH2, RP9, ZNRF4, SYT6, RORB, ITSN2, CD209D, BTK, ATP2B1, MAP3K6, MAP3K5, ZFP930, U2AF1, ZRSR1, CDH26, ZFP936, CLCA1, PLD6, RXRG, OLA1, POLB, NUDT10, OGFOD1, PRDM9, PITPNM3, HNF4A, ZGLP1, NEK9, ADAMTS1, UNC13B, FUS, ERMP1, GNAI2, ZCRB1, AKAP13, UBA5, PPAT, PLAGL1, THTPA, MACF1, ZMPSTE24, AEBP2, BMP1, TGFBR1, AGBL4, S100A11, TNNI3, AFG3L1, PPA2, ZFP609, ATP13A4, ZFP940, CBLC, ZFP442, RNF7, HSPB11, RAPSN, COX1, DZIP1L, ZFP947, TRAFD1, ZFP949, PRKRIR, ZMAT1, PPARA, SH3RF1, PXDN, ZFP46, PPARG, GLI3, STAC3, GM14391, NUBP1, GM14393,	386	18	2.39E-08	HCCS, GDA, ITSN1, CIAPIN1, BRPF1, DDAH1, CRYAB, NME6, OGFOD1, PITPNM1, NME3, ZNHIT1, PELO, ZNHIT3, TMEM129, RPS27, CDADC1, ZC3H13, NPLOC4, KLF13, MAP2K2, TGFBR2, GAS6, HPCA, MZF1, CDC42BPA, CDC42BPB, ZFP385A, RNF187, MMP24, FBXL19, USP19, STAC2, SIK3, USP13, CAR14, ZSWIM6, ADNP, MMP17, ZSWIM7, VPS8, GNAL, TAF15, KSR1, PHYHD1, REPS2, FHL2, NECAB3, NECAB2, PLA2G12A, ILVBL, LIMD2, TCEA2, FEN1, GALNT14, BRD1, PPP1R10, TRIM25, SDHB, MNAT1, LRP1, CDH15, PDE2A, SDHC, MCFD2, NARFL, DGKZ, GNA12, MIPEP, ZFP787, ZFP580, PVALB, PDE4A, ZFP688, CTDSP2, RPS27A, HMGCL, PHRF1, ZFP692, ACTN4, MTA3, PDXP, STIM1, ACTN2, RNASEH2A, KDM2A, ZFP791, TESK1, ZFP511,

			<p> P4HA3, NOS2,  COL11A2, IHH,  PRSS37, KCNMA1,  ZFP422, ACO1,  ZSWIM6, CYB5B,  RBBP6, PJA2,  RNF180, KSR2,  ZFP866, ASH1L,  GNAS, ZMYND11,  NKD2, ZFP712, TH,  FHL3, MYCBP2,  DGKA, DGKB,  MORC4, PPP2CA,  ETFDH, PRKAA2,  RBM26, LRSAM1,  ZFP57, ATP1A3,  ZFP119B, TRIM26,  ATP1A1, TRIM24,  SF3A2, CAPN2,  PCK1, ZFP9, ZFP106,  CYP4F18, SDHC,  ZFP207, CHN1,  NR5A1, CALM1,  APOBEC3, GM5136,  GNA11, LNX2,  LMO7, COX5B,  ZFP786, RNF141,  CYP2J11, TIMM9,  CTDSP2, ASPH,  2010111I01RIK,  AMY1, ZFP692,  KCND1, NUDT5,  PDE4D, PDE4C,  SUZ12, ZFP598,  PDE5A, VSNL1,  RNF138, COL1A1,  CAR4, PPP5C, CAR7,  ARFGAP1, CNBP,  ASTN2, RIOK1,  LIN28A, ITGAM,  RBX1, VDR, PDE6C,  ZFP760, RNF168,  TRAF7, B4GALT6,  RNF14, RNF13,  TAF3, WHSC1,  ITGA4, CACNA1S,  GM14326, TRIM56,  PHF3, GM14325, </p>			<p> ARFGAP1, ARFGAP2,  TRAF2, CNBP, ME3,  ELAC2, TIMP4, ZFP763,  TIMP3, ECE2, RNF165,  ADAM33, RNF169,  RNF167, DDX41,  B4GALT6, TRAF6, SDF4,  B4GALT7, TRAF4,  TRP53, B4GALT3,  B4GALT2, TRIP4,  EHMT1, AKAP8L,  MYL12B, RFNG, VAV1,  B3GAT3, EBF4, DOHH,  POLD1, CARS2, HIVEP1,  RPL37A, ABL2,  CACNA1A, HDAC7,  ZFP13, CYC1, ZKSCAN1,  NDUFS7, TRIM8,  HMOX1, ZCCHC9,  ZFPL1, E4F1,  CDK5RAP1, NDUFS2,  KDM5B, ZCCHC2,  ZFP553, TRPM7,  IRF2BP2, TOPORS,  ZFP652, PPM1F, ISCA2,  PPM1K, ZFAND2B,  CHFR, KDM6B, ACVR1,  CYP2U1, PHLPP1,  EGLN2, CABP5,  TIMM13, CDH2, CDH5,  DPF1, APLP1, CDH8,  NUDT8, RNF128,  NAIP5, GGPS1, ZFP524,  ZFP523, EHD1, CHD4,  VPS18, PDF, SLX1B,  CYP20A1, ADI1, CADPS,  RNF115, ZBED4,  SMPD1, ZFP536,  S100A3, S100A6, ALAD,  ZFP637, QPCTL,  ZFP335, SYT3, PRIM2,  NT5C2, CYHR1, S100A1,  YDJC, POLL, ZC3HC1,  RPP21, ZFP467,  NUDT11, PRDM8,  UHRF2, PIAS4, SIAH1A,  UBR5, DEAF1, MYO9B,  CALR, ZFP457, RIMS1, </p>
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			<p>SALL4, ZFP146, S100B, ITGA6, GM14322, FREM2, ZIC4, PRICKLE2, CYP2C38, AGTPBP1, MOB3C, PINK1, ZKSCAN1, DPH3, UQCRFS1, RHOU, TNIP2, IMPDH1, IMPDH2, ARL1, HMGCLL1, CYCS, RING1, MBNL2, COQ7, XRCC6BP1, ARL3, LAP3, RFWD2, ACVR2A, PPM1D, RFWD3, ZFP658, MDM2, UGP2, PEG3, AOC3, ING3, ZFP397, GALNT7, ADAMTS16, GLIS1, TIPARP, CYTB, BRSK1, NR3C1, ZFP661, CDH4, DROSHA, ADAT1, NPTX1, RASGRP1, ZFP664, PIKFYVE, TGM3, PLCD4, SCO1, ZBTB7B, INSM1, CYP2C70, WDFY1, ZFP386, CYP21A1, GM14410, TAB2, TAB3, SIRT3, GM14418, APIP, ALKBH3, ALKBH2, ALKBH5, ARAP1</p>			<p>TCF20, GALNS, NR1H3, ZFP346, KLF6, ZBTB48, IKZF4, CRIP1, OSGEP, SCD2, S100A16, HPCAL1, GDE1, SMYD2, ZFP445, ZFP746, U2AF1L4, ZBTB43, PPA1, CBLB, RNF7, ATP13A1, ATP2A3, RNF5, KLF2, PLA2G4B, TSNAX, ZMAT2, ZEB2, TRMT1, EFHD2, CGREF1, MBTD1, ELOF1, NT5M, NR2F6, ALOX12B, NEURL1B, NOS3, NEURL1A, AGAP3, ZBTB22, ZFP423, ZBTB20, ZCCHC10, SF1, HERC2, GMPR, RBBP6, ADO, ZFP865, RAB11FIP3, ZFP866, DNAJC24, ZFP869, ZFP961, THAP7, ZFP64, THAP4, ZFP414, THAP3, USP5, CETN2, ZFP316, PFAS, CYB561D2, CYB561D1, MORC3, TYW1, DTNB, ZFP410, BAZ2A, RBM22, MEX3B, ATP1A2, ATMIN, SOD2, BLVRA, PAPOLA, FYN, CYP4F16, ARSA, CALM2, APOBEC3, SLC6A1, CLYBL, FAHD1, GATA5, SLC25A23, TRIM44, RNF38, RABGEF1, PPP4C, ITPK1, RNF31, PICK1, TRIM41, ISCU, CPE, CHPF, MORC2A, RNF26, KALRN, ZC3H3, AMZ2, ITGB2, LMAN2, ZFAND1, RIOK2, RBX1, MYL9, PRR3, RNF10, STK38L, EWSR1, ACY3, ZMYM3, GALT, MUL1, UBOX5, TET2, LHPP,</p>
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							GM14326, GCM1, PRICKLE1, DNMT1, ASNA1, PHF8, GPATCH8, EPS15L1, CBFA2T3, PSPH, PDCD2, GSS, DGCR8, MT3, EGR1, SUOX, ELP3, WBSCR17, NRXN3, PFKP, ADIPOR2, ZFP322A, PRKCE, GZF1, NUCB1, MYT1L, DDR1, EYA1, DIS3L2, AOC2, ZFP367, FTL1, ZC4H2, NANOS1, GLRX2, RASGRP2, PLCD3, MUS81, SCNM1, PLCD1, ZFP292, CAMK2A, ENO1, DUS3L, DTX4, ZBTB7A, DTX1, SIRT4, SIRT7, PARK2, SPARC, HBA-A1, TDP2, ZC3H11A, ATP8A1, RNF41	
<b>Zinc-finger</b>	123	8	1.92E-04	PNMA3, DZIP1, LNX2, ZNRF4, RP9, RORB, ZFP786, BTK, RNF141, ZFP930, U2AF1, ZRSR1, ZFP936, ZFP692, PLD6, RXRG, SUZ12, PRDM9, ZFP598, HNF4A, ZGLP1, RNF138, UNC13B, ARFGAP1, FUS, CNBP, ZCRB1, AKAP13, LIN28A, RBX1, PLAGL1, VDR, ZFP760, RNF168, TRAF7, RNF14, RNF13, AEBP2, TAF3, WHSC1, ZFP609, ZFP940, GM14326, CBLC, TRIM56, PHF3, GM14325, ZFP442, SALL4, RNF7, ZFP146, RAPSIN,	187	9	1.07E-05	ZFP637, ZFP335, ZFP787, ZFP580, BRPF1, GATA5, TRIM44, RABGEF1, ZFP688, RNF38, CYHR1, RPS27A, RNF31, ZFP692, PHRF1, ZC3HC1, MTA3, TRIM41, ZFP467, PRDM8, UHRF2, PIAS4, KDM2A, UBR5, SIAH1A, ZFP791, MORC2A, ZNHIT1, RNF26, ZFP511, ZNHIT3, ZC3H3, ARFGAP1, DEAF1, TRAF2, ARFGAP2, CNBP, MYO9B, ZFP763, ZFAND1, RIMS1, ZFP457, RBX1, TMEM129, TCF20, RPS27, PRR3, RNF165, RNF169, RNF10, RNF167, TRAF6,

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								VPS18, DTX1, SLX1B, MEX3B, PPP1R10, TRIM25, PARK2, ATMIN, MNAT1, RNF115, ZBED4, DGKZ, ZC3H11A, ZFP536, RNF41
Zinc	153	10	8.77E-04	APOBEC3, PNMA3, GM5136, DZIP1, LNX2, LMO7, ZNRF4, RP9, RORB, COX5B, ZFP786, BTK, RNF141, ZFP930, TIMM9, U2AF1, ZRSR1, 2010111I01RIK, ZFP936, ZFP692, CLCA1, KCND1, PLD6, RXRG, SUZ12, PRDM9, ZFP598, HNF4A, PDE5A, ZGLP1, RNF138, ADAMTS1, CAR4, UNC13B, CAR7, FUS, ARFGAP1, ERMP1, CNBP, AKAP13, ZCRB1, UBA5, LIN28A, RBX1, PLAGL1, VDR, ZFP760, ZMPSTE24, RNF168, TRAF7, RNF14, RNF13, AEBP2, BMP1, TAF3, AGBL4, WHSC1, AFG3L1, ZFP609, ZFP940, GM14326, CBLC, TRIM56, PHF3, GM14325, ZFP442, RNF7, SALL4, ZFP146, S100B, RAPSN, GM14322, ZIC4, DZIP1L, PRICKLE2, ZFP947, ZFP949, TRAFD1, PRKRIR, ZMAT1, SH3RF1, PPARA, ZFP46, AGTPBP1,	237	12	3.55E-05	S100A3, ALAD, GDA, ZFP637, QPCTL, ZFP335, BRPF1, DDAH1, CYHR1, S100A1, ZC3HC1, RPP21, CRYAB, ZFP467, PRDM8, UHRF2, PIAS4, UBR5, SIAH1A, ZNHIT1, ZNHIT3, DEAF1, MYO9B, CALR, ZFP457, RIMS1, TMEM129, TCF20, RPS27, CDADC1, NR1H3, ZFP346, IKZF4, ZC3H13, KLF6, ZBTB48, NPLOC4, CRIP1, KLF13, SMYD2, ZFP445, U2AF1L4, ZFP746, ZBTB43, CBLB, RNF7, RNF5, MZF1, CDC42BPA, KLF2, CDC42BPB, ZFP385A, ZMAT2, RNF187, ZEB2, TRMT1, MMP24, FBXL19, USP19, STAC2, MBTD1, ELOF1, NR2F6, NOS3, NEURL1B, NEURL1A, AGAP3, USP13, ZBTB22, ZFP423, ZCCHC10, ZBTB20, CAR14, SF1, ZSWIM6, ADNP, MMP17, ZSWIM7, HERC2, RBBP6, VPS8, ZFP865, ZFP866, TAF15, DNAJC24, ZFP869, KSR1, ZFP961, THAP7, ZFP64, THAP4, THAP3, ZFP414, USP5, FHL2, ZFP316, MORC3, DTNB, ZFP410, LIMD2, TCEA2, BAZ2A, RBM22,

				<p> <b>MOB3C, PPARG, ZKSCAN1, DPH3, GLI3, STAC3, GM14391, GM14393, NOS2, TNIP2, IHH, ZFP422, ZSWIM6, RING1, MBNL2, RBBP6, LAP3, RFWD2, RNF180, PJA2, KSR2, RFWD3, ZFP866, ZFP658, ASH1L, MDM2, PEG3, ZMYND11, ING3, ZFP397, ADAMTS16, GLIS1, ZFP712, TIPARP, FHL3, ZFP661, NR3C1, MYCBP2, DROSHA, DGKA, ADAT1, DGKB, MORC4, RASGRP1, PIKFYVE, ZFP664, RBM26, ZBTB7B, INSM1, LRSAM1, WDFY1, ZFP57, ZFP386, ZFP119B, TRIM26, GM14410, TRIM24, SF3A2, TAB2, TAB3, SIRT3, GM14418, ZFP9, ZFP106, CHN1, ZFP207, APIP, ARAP1, NR5A1</b> </p>				<p> <b>BRD1, MEX3B, PPP1R10, TRIM25, ATMIN, BLVRA, MNAT1, DGKZ, APOBEC3, MIPEP, ZFP787, ZFP580, GATA5, TRIM44, RABGEF1, ZFP688, RNF38, RPS27A, RNF31, ZFP692, PHRF1, PICK1, MTA3, TRIM41, CPE, KDM2A, ZFP791, MORC2A, RNF26, ZFP511, ZC3H3, ARFGAP1, TRAF2, ARFGAP2, CNBP, ELAC2, AMZ2, TIMP4, ZFP763, TIMP3, ZFAND1, RBX1, ECE2, PRR3, SLC30A1, RNF165, RNF169, ADAM33, RNF10, SLC30A3, RNF167, TRAF6, DDX41, TRAF4, EWSR1, TRP53, EHMT1, TRIP4, ACY3, ZMYM3, AKAP8L, GALT, MUL1, UBOX5, TET2, VAV1, GM14326, GCM1, EBF4, PRICKLE1, POLD1, DNMT1, CARS2, HIVEP1, RPL37A, ASNA1, PHF8, GPATCH8, HDAC7, ZFP13, ZKSCAN1, CBFA2T3, PDCD2, TRIM8, ZFPL1, ZCCHC9, E4F1, GPC1, KDM5B, MT3, EGR1, ZCCHC2, ZFP553, TRPM7, ADIPOR2, IRF2BP2, ZFP322A, TOPORS, PRKCE, GZF1, ZFP652, MYT1L, ZFAND2B, CHFR, KDM6B, ZFP367, ZC4H2, NANOS1, TIMM13, APLP1, DPF1, RASGRP2, NAIP5,</b> </p>
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								RNF128, ZFP524, ZFP523, SCNM1, ZFP292, CHD4, DUS3L, DTX4, ZBTB7A, VPS18, DTX1, SLX1B, SIRT4, SIRT7, PARK2, RNF115, ZBED4, SMPD1, ZC3H11A, ZFP536, RNF41
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<sup>1</sup>Huang, D. W.; Sherman, B. T.; Lempicki, R. A. Systematic and integrative analysis of large gene lists using DAVID Bioinformatics Resources. *Nature Protoc.* **2009**, *4*, 44-57.