

SUPPLEMENTARY ONLINE DATA

Reprogramming of pancreatic exocrine cells towards a beta cell character using *Pdx1*, *Ngn3* and *MafA*Ersin AKINCI, Anannya BANGA, Lucas V. GREDER, James R. DUTTON and Jonathan M. W. SLACK¹

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Pdx1:

AGAGATCAGCCTGCTGAGAGAGAAATTGAAA(CAAGTGCAGGTG)TTCGCGGGGCTGGC
CTCC(TTCTTAAGG)CAGGGCCAGG(CCAAT)GGTGGCCCCAGGCTGAA(CCACGTG)GGGTG
CTCAGAGCCTATGGCACGGCAGCGCTCTGTCTCGCCAGCCTGTGGTCCCAGG
(GAGCA)GTGGAGAACTGTCAAAGCAGTCAGGGTGCGCTGAGAGTCCGTGAGCTGCCAG
CGCCTAAGCCTGGCTTAGCTCCCTACCCGGCTGCCGGCCCCGAAGTGCCGGCTGCC
ACCATGAATAGTGAGGAGCAGTACAGCGGCCACACAGCTCTACAAGGACCCGTGC
CATTCCAGAGGGTCCGGTCCAGAGTCTAGTCTAATCCCTCGTGTACATGGCCG
CCAGCCCCCACCTCCGCCACCCAGTTGCAGGCTCGCTGGGAACGCTGGAACAGGGA
AGTCCCCGGACATCTCCCATACGAAGTGGGGCCGCTGCCGTGACCCGGCTGGCGC
ACCTCCACCAACCTCCAGCTCAGCTGGCTGCCCATCACCTCCGGACCTTCCC
ATGGAACCGAGACTGGGGCTGGAAGAGCCCAGCCGTTACCTCCCTTCCCAG
TGAAATCCACCAAGCTCACGCGTGGAAAGCCAGTGGCAG

Ins1:

ATAATC(TAATT)CCCTAGGTCTAAGTAGAGTTGT(TGACGT(C)CAAT)GAGCGCTTCTGC
AGACTTAGCACTAGGCAAGTGT(TGAAAT)(TACAGCTTCAGCC)CCTCTCG(CATCTG)CC
TACCTACCCCTCCTAGAGCCC(TTAAT)GGGCAA(ACGGCAAAGTCC)AGGGGGCAGAGAGG
AGGTGCTTGGAC(TATAA)AGCTAGTGGAGACCAGTAACCTCCAACCTAAGTGAC(CAGC)
ACAATCATAGACCATCAGCAAGCAGGTATGTACTCTCTGGGTAGGCCGTTCCCCAGCC
AAAACCTAGGACTTTAGGAAGGATGTGGTCTCTTACATGGACCTTCTAGCCT
AACCCCTGCTATCTCAGGTCAATTGTCACATGGCCCTGTGGATGCGCTTCTGCCCTG
CTGGCCCTGCTCGTCCCTGGGAGGCCAACGCTGCCAGGCTTTGTCAAACAGCACCTTG
TGGTCTCACCCTGGTGGAGGCTCTGACTCTGGTGTGGGAACGTTTCTTACACAC
CCAAGTCCCCTGCTGAAGTGGAGGACCCGCAAGTGCACAACTGGAGCTGGTGGAGG
CGGAGGCCGGGATCTCAGACCTTGGACTGGAGGTTGCCGGAGAAGCGTGGCATTG
GGATCACTGCTGCACCATCTGCTCCCTACCAACTGGAGAAACTGCAACTGCAACTG

Ins2:

AGCAAGGCCCTCTGGCTCTCTATGGCTCTGAGGAGACCTCTCCACACTGCCCTGGCTTCCC
ACCCCTGGTCTTACCTGAGCTCTTACCTGCTCTCTTGCATTCACTGATCTCTGCTACAG
TAGGTCCACGGTGGTCTCCAGATAACCGGAGTGTGAGTGGCTGCAGCACTTCTGGGGGA
CAAGAAGTAGGGAGCAAGGGGCTCACAGTCAAGTCTGGTGTATAAAGCCCTGCATAGG
GTAGAGTTCTCGCTCATGCAACGACACCAAGGGTTTGTCTGCTGCTCGGGGAACAGGGC
AGTACCAAAATCAGGAACAGAAAGAGTCAGGATCCCCAACCTCCAAGTGGAGGCTGAG
AAAGGTTTGTAGTGGTAGAGTATGTAAGAGATGGAGACAGTGGCTGAGCTCTG
AAGCAAGCACCTTATGGAGAGTGTGACCTTCAGGTGCAAATCTAAGATACTACAGGAG
AATAACCATGGGCTTCAGCCCAGTCTGACTCCGAGTGGGATATGGTTGTGGAAGGAG
AGATAGAAGAGAAGGGACCTTCTTGAATTCTGCTTCTACCTCTGAGGGTGAGCT
GGGGTCTCAGCTGAGGTGAGGACACAGCTACAGTGGAACTGTGAAACAACAGTTCAAGG
GACAAAGTTACTAGGTCCCCAACAACTGCAAGCCTCTGGGAATGATGTGGAAAATGCT
CAGCCAAGGACAAGAAGGCCCTACCCCTCTGAGACAAATGTCCTCTGCTGTAACGGTC
ATCAGGCCACCCAGGAGCCCTATTAAGACTC(TAATT)CCCTAAGGCTAAGTAGAGGTGT
(TGTGT(CC)AAT)GAGCACTTCTGCAAGACCTAGCACCAGGCAAGTGT(TGAAAC)(TGCAG
CTTCAGCC)CCTCTGG(CATCTG)CTGATCCACCC(TTAAT)GGGACAAACAGCAAAGTCCAG
(GGGTCAAGGGGGGGGGTG)TTTGGAC(TATAAA)GCTAGTGGGGATTCAAGTAACCCCCAGCC
TAAGTGACCAAGCTACAGTCGGAACCATCAGCAAGCAGGTATGTACTCTCCAGGGTGGGCC
TGGCTTCCCCAGTCAAGACTCCAGGGATTGAGGGACCGCTGGGCTTCTTACATGTA
CCTTCTGCTAGCCTCAACCTGACTATCTCAGGTATGTTCAAACATGGCCCTGTGGATC
CGCTTCTGCCCCCTGTCGGCCCTGCTCATCTCTGGAGCCCCGCCCTGCCAGGCTTGTG
AAACAGCACCTTGTGGTCTACTGGTGGAGCTCTACCTGGTGTGAGCCTGG
ATTCTCTACACACCCATGTCCCCGCCGAAGTGGAGGACCCACAAG

Figure S1 Location of the primer pairs used for ChIP, and CpGs sequenced for methylation analysis, in the genomic sequences of *Pdx1*, *Ins1* and *Ins2*

Black, proximal promoter; blue, 5' untranslated region; red, first exon. The sequences in parentheses indicate known regulatory elements. Primers for ChIP qPCR are highlighted in yellow. CGs shown in bold type were sequenced in the DNA methylation assay.

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Table S1 Primer pairs used in RT-PCR, ChIP qPCR, DNA methylation analysis and qRT-PCR

Primer name	Forward sequence (5'-3')	Reverse sequence (5'-3')
Primers used in RT-PCR		
Ins1	CTACAATCATAGACCATTAGCA	CAGTTGGTAGAGGGAGCAGAT
Ins2	CCCTGTGGATCGCTTCCCTGC	GTCACAAAGGCTGAAGGTCA
Gcg	GCTGGCAGCATCCCCCTCAA	CCTTGCTGCCCTGGCCCTCC
Ppy	CTATCCACTTGGGGCTCT	ATCAAACCCACAGAAGGC
Sst	CCCAGACTCCGTAGTTCT	GAAGTCTTGAGCCAGCTT
Iapp	GGCTGTAGTCTCTGAAGCTT	AAGGTTGTTGCTGGAGCGAA
Pax4	TGGAGAAAGAGTCCAGCGT	CTTGGGATGTTGTCAGTG
Neurod1	TTTCAAAACACGAACCATCCA	CCTGTTCTTCAAAGGCAG
Nkx2.2	CACCGCGCTAACAGCCAGGA	GGAGAGAGCGGTGGGGCAGA
Arx	AGGACGCCAGGGCAAGGAT	TCTCCCCTTGCGCAGCTTG
Pdx1	GTAGTAGCGGGACAAAGGAC	CAATTGGGAGCCTGATTCTC
Ngn3	AAACTTCAAGGGAGCAGAG	CATCCCTGAGGTTGGAAAAA
MafA	TCCAGGCTGGTGCAGCGAAG	GCAAGCCCACCTCAGGAGCG
Isl1	TGATGCGCAGCCGCTCTAA	TGACTCGGGACTGAGGCCG
Hlx9	CCTGTCGCGACCCAAGCGTT	CCCAGCAGCTCTCTCCGT
Pax6	GGCCGTGCGACATTCCCGA	GCGCTCTGCCCTACTGTT
Hes1	GGCCACCTGGCAACTGCAT	GCTGGAAGGCGACACTGCG
Glut2	GCCATCTTCTCTCGTCAG	ACCTGGTTCCCTCTGGTCT
Kir6.2	CTGGAAGGAAGGAGCTTGT	CAGTGTCCCCAGACAAAGT
Sur1	AGCTGCGCTCTGCCTCACG	GGCACCCCTGCTGGCTCTG
Amylase	GCCTACTGACAGAGCCCTG	TGGTCCATCCAGTCATTCA
Ptf1a	CCTCTCAAAGGTAGACACGC	CTTGGGATGTTGTCAGTG
Cbe	GACCGGGCGTACAGTCGCG	CTGCGCCCCACCGTGTAGAT
Pcsk2	ACACAGCTCCGACATTGCA	TGAGATCCACAACGCCCTCCA
Prss1	GTGTATCCTCAACGATCTGT	CACTCTGATCCTAGCCCCTTG
Cpa1	CCAGAAGTCCAACGCAAGT	CAGTGTGGCAATGAGACT
Ctrb1	GAATAGCATCTCCCGTTGAC	GTCCTGCTTGCCTTGT
Rbpj1	CATCTCGAACACACCTTG	CTCCAGTGCCTCATATCAGC
Beta Actin	TCCGTAAAGACCTCTATGCC	GGAGGGGCCGGAACATCGT
mNgn3	CCGGATGACGCCAACTTACA	ACACCAGTGTCCCCGGAG
mPdx1	CCGGACATCTCCCATACGAAGT	CGCACAACTTGTCCGGCTCT
mMafA	ATCATCACTCTGCCACCAT	AGTCGGATGACCTCTCCTT
Primers used in ChIP qPCR		
Ins1	CCAATGAGCGCTTCTGCAGACTT	AGGAGAGTACATACTGCTGCT
Ins2	ACCAGGCAAGTGTGGAAACTGC	ATGTAAGAGAACGCCACAGCGT
Pdx1	AGAGATCAGCCTGCTGAGAGAGAA	TACTGCTCCTCACTATTGATGGTGGC
Gapdh	CTTACGGGTGACGTAGCTCA	TTTCACCTGGCACTGCACAAGAAG
Primers used in DNA methylation assay		
Ins1 1st set	TTTAAAGTTAACAGAGTTGTA	CCCTAAATTAACTAAAAAAC
Ins1 2nd set	TTTTTTGTTTTGTTGGTTTTGTT	CACCCAACTCCAATTAAACACTTAC
Ins1 3rd set	TTTTGGGAGTTAACAGTTGTTAG	ATACAACACTAATCCACAAATACCAC
Ins2 1st set	TTTGAGTTTTTATTTGTTTTT	CTAATTAAATACCTCTTATTC
Ins2 2nd set	GATTATAAAAGTTAGTGGGATTAGTAATT	TATTAAACAAAACCTAACAAAC
Ins2 3rd set	GTATTTTGTTGGTTTATTGGTG	ACTTACCTTATAAAATCCTCACTTC
Pdx1 1st set	AGGAGAGATTAGTTGTTGAGAGAGAA	AAAAACTACAAACCAAACCTAAAC
Pdx1 2nd set	ATTATGAATAGTGGAGGAGTAGTATT	AAAAACTTCCCTATTCAAC
Pdx1 3rd set	GTTGAATAGGGAAAGTTTT	ACTTACCTACCCACTAACCTTCCAC
Primers used in qPCR		
Ins1	CAATCATAGACCATTAGCAAGC	AGAAACACGTTCCCCAC
Ins2	CCCAGGCTTTGTCAAACAG	GTGCCAAGGCTGAAGGTC
Iapp	CCACTGAAAGGGATCTGAGAC	TCCTGTTGTCACCTGAG
Pax4	GGGCAGTATCCGATTCTGGT	GGCATCTGTGTTCCATTG
Neurod1	ATGTCTTCCACGTCAGCC	GAGAAGTGGCATTGATGCTG
Nkx2.2	GGTCAGATCTGGTCCAAAAC	GTCACCTCCACCTTCTCAG
Arx	ATCTAACCCATCCCCAACAC	CTCTTCTGGTACTGATGCTC
Endogenous Pdx1	CCCGAGCTTCTGAAACTTG	CTTTCTATTGTCCTCAGTTGG
Endogenous Ngn3	TCCAGACGCAATTACTCCAG	CTAGTCTCCGGGCTAAAG
Endogenous MafA	TCTTCTGTGAGCGCG	TCAGACTCGAACCGAGG
Isl1	TTGTTAGGGACGGGAAAC	CTACACAGCGAACATTG
Hlx9	GCTTCTACTCTGTATCTCG	TTCCCAAAGAGGTCATG
Hes1	AGAAAAATTCTCGTCCCCG	TTTCATTATTCTGCCGGC
Glut2	CACATCCACTTGGCTATCTG	TCAGTGCCTTACTGCTTT
Kir6.2	TCAGTAAAGCAATGAGCAGGG	CAACCTCTGGACTGATATGCC
Sur1	AGAAGCTCTAGACTACCG	TGAGGAGTTGGAGATGGAG
Amylase	GCAGACCTTCTTCAAGAG	ACAAAACCCAAACCTTCTCC
Ptf1a	AGGACCCCGAAGAAACTCAAC	CAATATGCAACAGACACAGCC
Cbe	ATGGTAATGAGGCCGGTGG	AGGGCATGATGATGGATTG

Table S1 Continued

Primer name	Forward sequence (5'-3')	Reverse sequence (5'-3')
Pcsk1	CTCAGCCCTCTACTTGTG	CATTGACAAACTGCCCTTCG
Pcsk2	GCATAAAGACGGAGAGGAAGAG	TGGAAAAGTGGTACAGGCC
Prss1	GTTATCCTCCAACGATCTTG	CACTCTGATCCTAGCCCTTG
Cpa1	CCAGAAGTCCAAC TGCAAGT	CAGTCTGTGGCAATGAGAACT
Cpb1	CACGTTGCTTATCGTACCTCA	GCCCTCTCACTACAGTTGACTT
Ctrb1	GAATAGCATCCTCCGGTGTAC	GTCCTGCTTGCCCTTG
Rbpj1	CATCTCCGAACCACACCTG	CTCCAGTGCCCTATATCAGC
Gapdh	TCCAGTATGACTCTACCCACG	CACGACATACTCAGCACCGAG

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