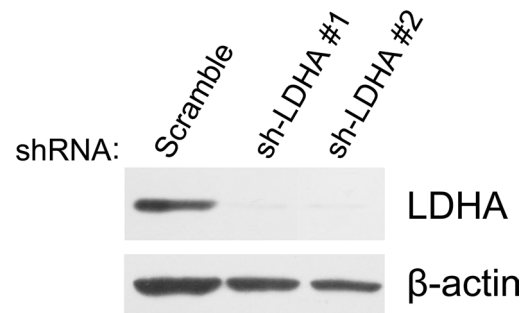
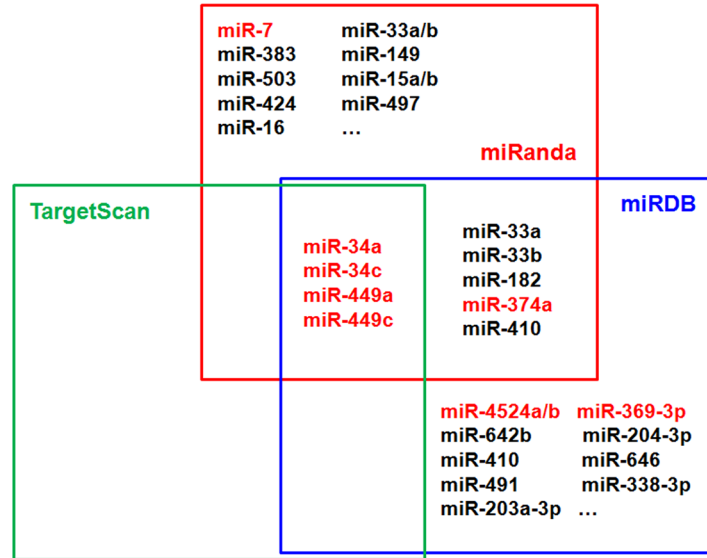


SUPPLEMENTARY FIGURES AND TABLES

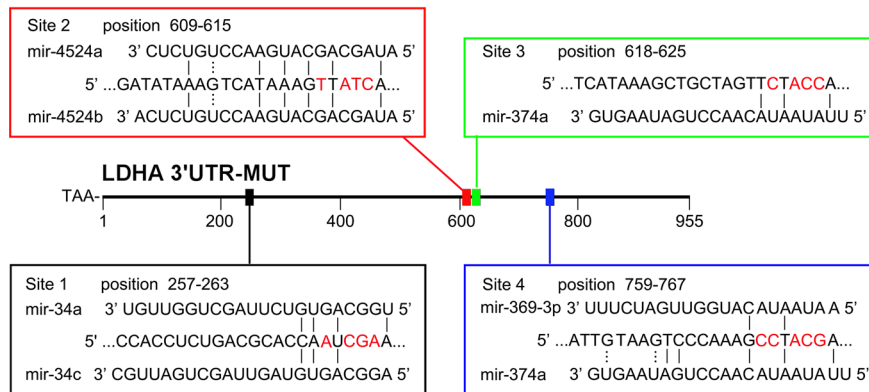


Supplementary Figure 1: Western blot analysis of LDHA in cells stably expressing LDHA-shRNA or scramble-shRNA.

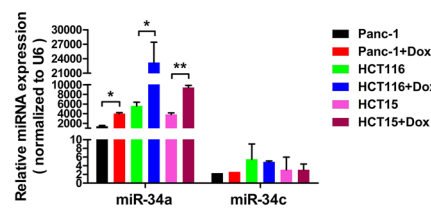
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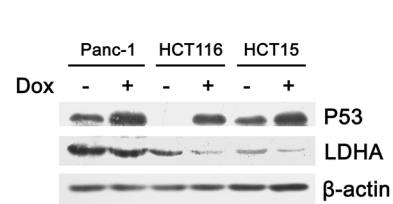
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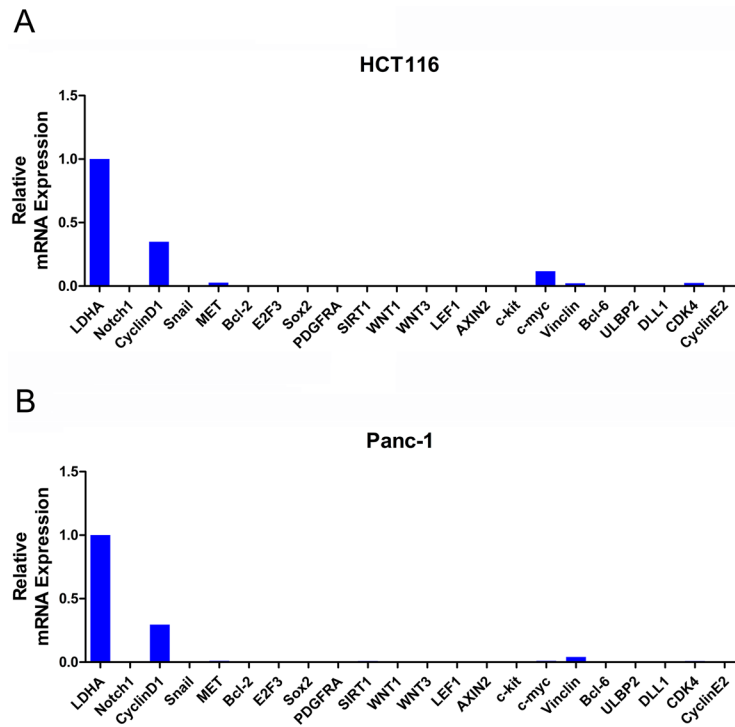
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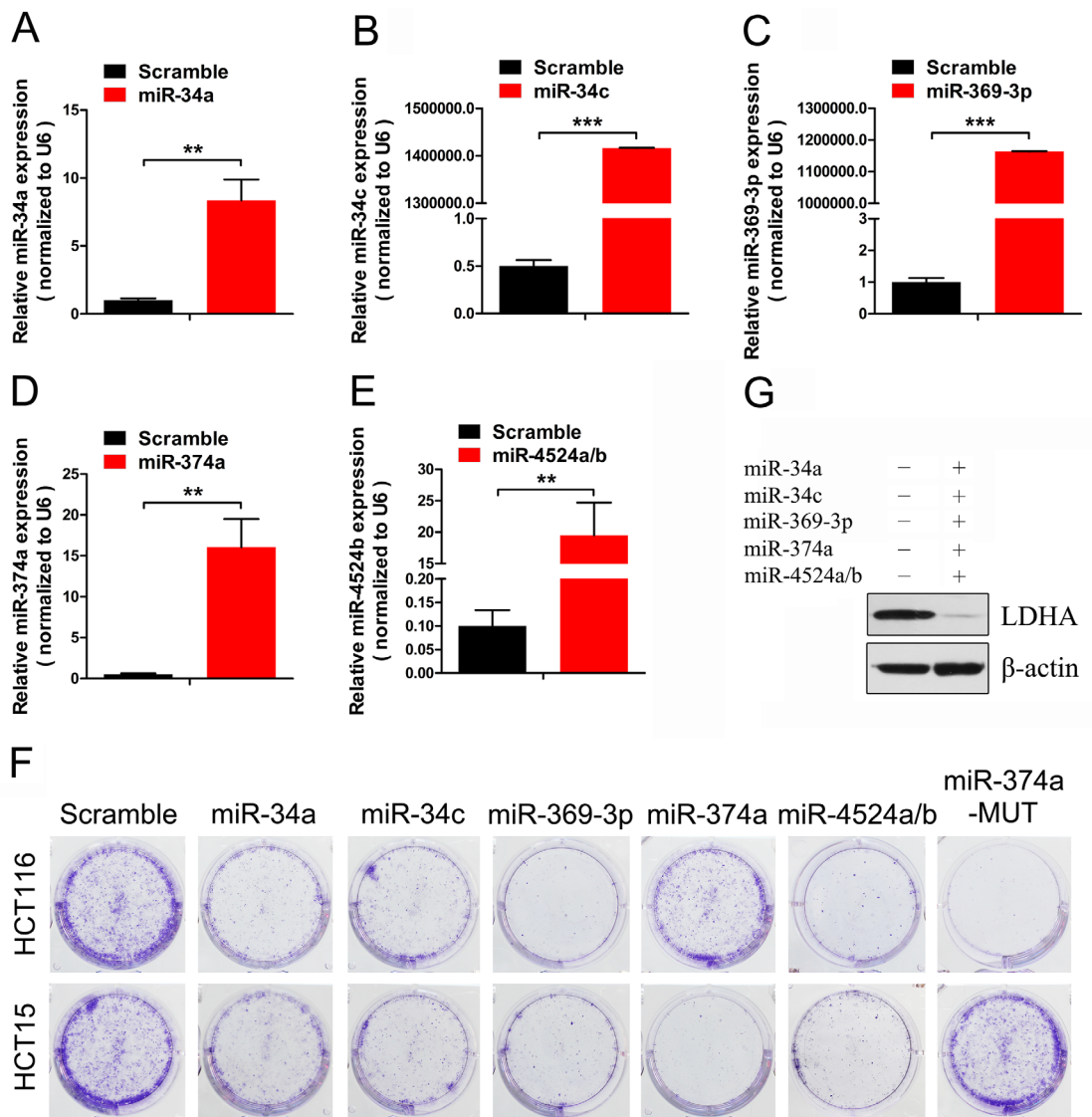
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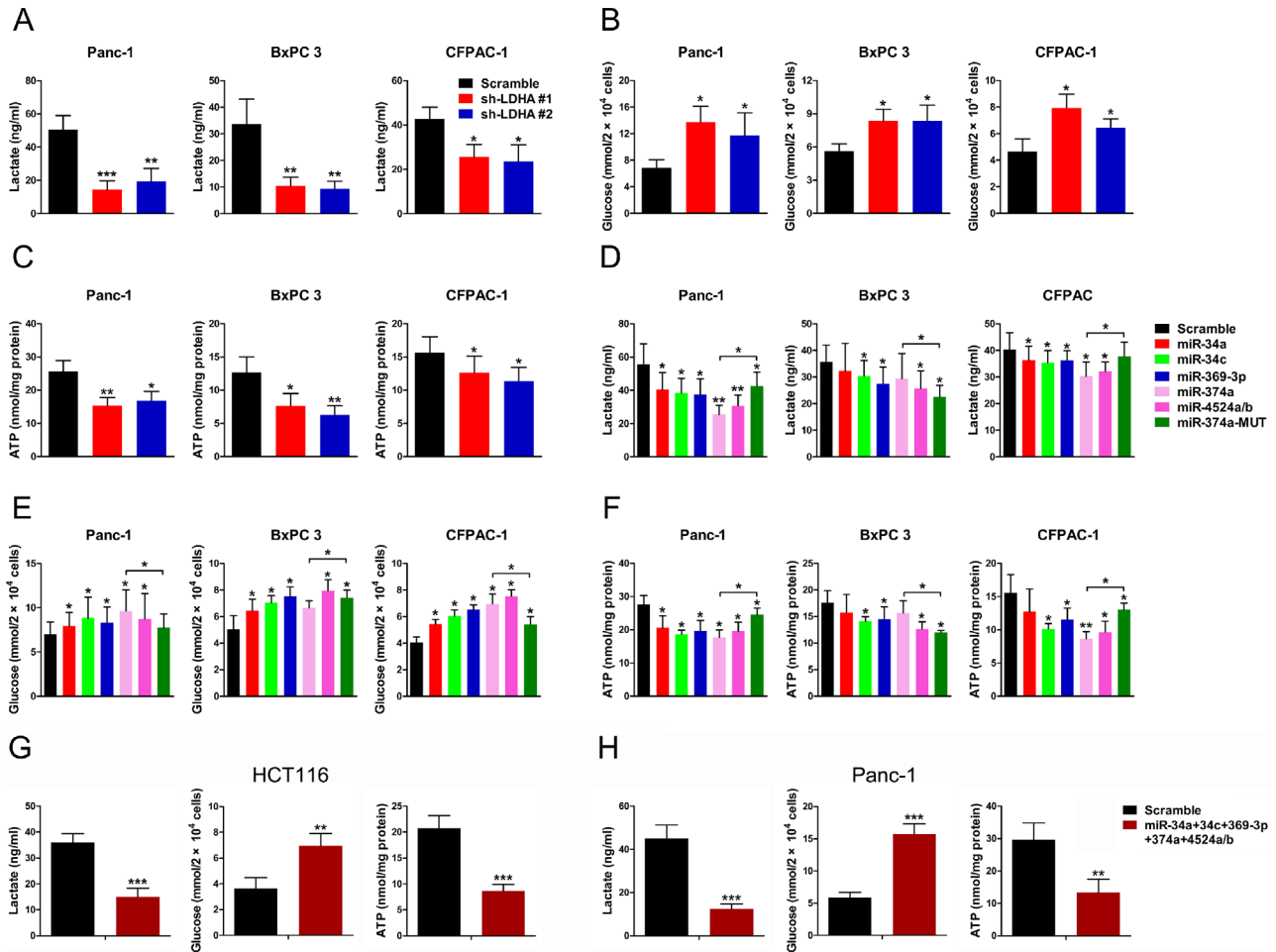
Supplementary Figure 2: (A) miRNAs, predicted by miRanda, TargetScan and miRDB, target LDHA. miRNAs that were predicted by two or more algorithms are indicated. (B) Sequences of indicated miRNAs and nucleotides mutated in LDHA-3' UTR-MUT. (C) RT-PCR analysis of miR-34a and miR-34c in Panc 1, HCT116, and HCT15 cells treated with doxorubicin. (D) Western blot analysis of P53 and LDHA in Panc 1, HCT116, and HCT15 cells treated with doxorubicin.



Supplementary Figure 3: (A, B) RT-PCR analysis of the target genes of miR-34 in HCT116 (A) or Panc-1 (B) cells.



Supplementary Figure 4: (A–E) RT-PCR analysis of miR-34a (A), miR-34c (B), miR-369–3p (C), miR-374a (D), and miR-4524a/b (E) in cells stably expressing indicated miRNA. (F) Colony formation assay results. Five hundred HCT116 or HCT15 cells stably expressing indicated miRNAs or scramble cells were plated in 6-well plates supplemented with 10% FBS for 2 days followed with 2% FBS for a further 12 days. Colonies were stained with crystal violet. The number of colonies was counted from three independent experimental wells. (G) Western blot analysis of LDHA in HCT15 cells stably expressing indicated miRNAs.



Supplementary Figure 5: (A, B, C) Lactate (in the culture media) (A), Glucose (in the culture media) (B), and intracellular ATP levels (C) in human pancreatic cancer cell lines stably expressing LDHA-shRNA and scramble-shRNA. (D, E, F) Lactate (in the culture media) (D), Glucose (in the culture media) (E), and intracellular ATP levels (F) in human pancreatic cancer cell lines stably expressing indicated miRNAs or scramble cells lines. (G, H) Analysis of Lactate (in the culture media), Glucose (in the culture media), and intracellular ATP levels in HCT116 (G) or Panc-1 (H) cells stably expressing miR-34a, miR-34c, miR-369-3p, miR-374a, and miR-4524a/b compared with scramble cells.

Supplementary Table 1: The sequence in position 618–625 of LDHA 3' UTR

| Cell lines | Sequence |
|------------|----------|
| HCT116 | TATTGAT |
| HCT15 | TATTATA |
| HT29 | TATTATA |
| Panc-1 | TATTATA |
| BxPC 3 | TATTGAT |
| CFPAC-1 | TATTATA |

UTR: untranslated region

Supplementary Table 2: List of primer

| Name of primer | Sequences of primer |
|----------------------|--------------------------------------------------------------------|
| RT-LDHA-F | TTGACCTACGTGGCTTGAAG |
| RT-LDHA-R | GGTAACGGAATCGGGCTGAAT |
| 3'-UTR-LDHA-F | GCGTTTAAACCCTGTCTAGGCTACAACAGGAT |
| 3'-UTR-LDHA-R | CCCTCGAGTTGTTACCAGCTTCCAGAGGA |
| Pri-miR-34a-F | GCTCTAGATTCCGCACCCACGAGCAG |
| Pri-miR-34a-R | CGGAATTCCTTCTTATCAACAGGTGCTGG |
| Pri-miR-34c-F | GCTCTAGATCACTGTGTCTATTTGCCATCGTC |
| Pri-miR-34c-R | ATTTGCGGCCGCATAGGCAATGGATGAAAT |
| Pri-miR-369-3p-F | GCGAATTCTTCCACCAGGGAGGCCGTC |
| Pri-miR-369-3p-R | ATTTGCGGCCGCCAAGGTGAGATTTGATACT |
| Pri-miR-374a-F | GGTCTAGAAGCTGACAATCCAGAGCCTGT |
| Pri-miR-374a-R | CGGAATTCGGGTTAGGGAAAGCCTGAATAT |
| Pri-miR-449a/c-F | GCTCTAGAGCACCATATAAACTGTGCATTG |
| Pri-miR-449a/c-R | GCGAATTCGTTTTTGGACTAGATTC AAGCAAAT |
| Pri-miR-4524a/b-F | GCTCTAGAGCAACACAATGGCATTAGGAAGC |
| Pri-miR-4524a/b-R | ATTTGCGGCCGCTTTATAACAGGTCTGCTGCAC |
| Flag-LDHA-F | CCGGAATTCGCCACCATGGATTACAAGGATGACG ACGATAAGGCAACTCTAAAGGATCAGCT |
| Flag-LDHA-R | CGCGGATCCTTGTTACCAGCTTCCAGAGGAC |
| miR-34a stem loop | GTCGTATCCAGTGCAGGGTCCGAGGTATTTCGC ACTGGATACGACACAACC |
| RT- miR-34a-F | GGTCCTGGCAGTGTCTTA |
| miR-34c stem loop | GTCGTATCCAGTGCAGGGTCCGAGGTATTTCGC ACTGGATACGACGCAATC |
| RT- miR-34c-F | GGTCCAGGCAGTGTAGTTAGCT |
| miR-369-3p stem loop | GTCGTATCCAGTGCAGGGTCCGAGGTATTTCGCAC TGGATACGACAAAGAT |
| RT- miR-369-3p-F | GGTCCAATAATACATGGTTG |
| miR-374a stem loop | GTCGTATCCAGTGCAGGGTCCGAGGTATTTCGCAC TGGATACGACCACTTA |
| RT- miR-374a-F | GGTCCTTATAATACAACCTGA |
| miR-4524a stem loop | GTCGTATCCAGTGCAGGGTCCGAGGTATTTCGCAC TGGATACGACTGAGAC |
| RT- miR-4524a-F | GGTCCATAGCAGCATGAACCT |
| miR-4524b stem loop | GTCGTATCCAGTGCAGGGTCCGAGGTATTTCG CACTGGATACGACGAGACA |
| RT- miR-4524b-F | GGTCCATAGCAGCATAAGCC |

(Continued)

| Name of primer | Sequences of primer |
|----------------|-------------------------|
| RT-miRNA-R | GTGCAGGGTCCGAGGT |
| RT-Notch1-F | TGGACCAGATTGGGGAGTTC |
| RT-Notch1-R | GCACACTCGTCTGTGTTGAC |
| RT-CyclinD1-F | CAATGACCCCGCACGATTTC |
| RT-CyclinD1-R | CATGGAGGGCGGATTGGAA |
| RT-Snail-F | ACTGCAACAAGGAATACCTCAG |
| RT-Snail-R | GCACTGGTACTTCTTGACATCTG |
| RT-Met-F | GGTTCACTGCATATTCTCCCC |
| RT-Met-R | ACCATCTTTCGTTTCCTTTAGCC |
| RT-Bcl 2-F | GGTGGGGTCATGTGTGTGG |
| RT-Bcl 2-R | CGGTTCAAGTACTCAGTCATCC |
| RT-E2F3-F | GTATGATACGTCTCTTGGTCTGC |
| RT-E2F3-R | CAAATCCAATACCCCATCGGG |
| RT-Sox2-F | TACAGCATGTCCTACTCGCAG |
| RT-Sox2-R | GAGGAAGAGGTAACCACAGGG |
| RT-PDGFR-A-F | TTGAAGGCAGGCACATTTACA |
| RT-PDGFR-A-R | GCGACAAGGTATAATGGCAGAAT |
| RT-SIRT1-F | AGCCAATTCTTTTGTGTTTCGTG |
| RT-SIRT1-R | CGGCGTTTATCTTCGCTATCA |
| RT-WNT1-F | CGGCGTTTATCTTCGCTATCA |
| RT-WNT1-R | GCAGGATTCGATGGAACCTTCT |
| RT-WNT3-F | AGGGCACCTCCACCATTG |
| RT-WNT3-R | GACACTAACACGCCGAAGTCA |
| RT-LEF1-F | TGCCAAATATGAATAACGACCCA |
| RT-LEF1-R | GAGAAAAGTGCTCGTCACTGT |
| RT-AXIN2-F | TACACTCCTTATTGGGCGATCA |
| RT-AXIN2-R | TTGGCTACTCGTAAAGTTTGGT |
| RT-c-Kit-F | ACTTGAGGTTTATTCCTGACCCC |
| RT-c-Kit-R | GCAGACAGAGCCGATGGTAG |
| RT-c-myc-F | GTCAAGAGGCGAACACACAAC |
| RT-c-myc-R | TTGGACGGACAGGATGTATGC |
| RT-Vinculin-F | ATGGGTCAAGGGGCATCCT |
| RT-Vinculin-R | GGCCCAAGATTCTTTGTGTAAGT |
| RT-Bcl 6-F | ACACATCTCGGCTCAATTTGC |
| RT-Bcl 6-R | AGTGTCCACAACATGCTCCAT |
| RT-ULBP2-F | GTGGTGGACATACTTACAGAGC |

(Continued)

| Name of primer | Sequences of primer |
|----------------|--------------------------|
| RT-ULBP2-R | CTGCCCATCGAAACTGAACTG |
| RT-DLL1-F | TGTGACGAGTGTATCCGCTAT |
| RT-DLL1-R | GTGTGCAGTAGTTCAGGTCCT |
| RT-CDK4-F | CTGGTGTTTGAGCATGTAGACC |
| RT-CDK4-R | GATCCTTGATCGTTTCGGCTG |
| RT- CyclinE2-F | TCAAGACGAAGTAGCCGTTTAC |
| RT- CyclinE2-R | TGACATCCTGGGTAGTTTTTCCTC |