Figure S1. Database search analysis of GO 'Biological process' category on Uniprot. HDAC2 was found to be associated with 'histone H3 deacetylation' and histone acetylation sites (H3K14) regulated by HDAC2. HDAC2, histone deacetylase 2; GO, Gene Ontology.

## GO-Biological process i

- behavioral response to ethanol 

  Source: RGD → cardiac muscle cell development 

  Source: RGD cardiac muscle hypertrophy Source: RGD cellular response to dopamine Source: RGD cellular response to heat Source: RGD > Source: RGD cellular response to hydrogen peroxide ■ cellular response to retinoic acid Source: RGD → cellular response to transforming growth factor deta stimulus Source: RGD cellular response to trichostatin A Source: RGD circadian regulation of gene expression Source: RGD dendrite development Source: RGD embryonic digit morphogenesis Source: RGD epidermal cell differentiation Source: RGD eyelid development in camera-type eye Source: RGD fungiform papilla formation Source: RGD hair follicle placode formation Source: RGD hippocampus development Source: RGD histone deacetylation Source: RGD
- histone H3 deacetylation
   Source: RGD
   histone H3-K27 methylation
   Source: RGD
   histone H4 deacetylation
   Source: RGD

## Child Terms This table lists all terms that are direct descendants (child terms) of GO:0070932 Child Term Relationship to GO:0070932 is\_a GO:1990619 histone H3-K9 deacetylation is\_a GO:0071572 histone H3-K56 deacetylation GO:0031078 histone deacetylase activity (H3-K14 specific) part\_of GO:1990596 histone H3-K4 deacetylation is\_a

Figure S2. Prediction of c-Myc binding sites at the miR-30a-5p gene promoter region. Location of the predicted c-Myc binding sites at the miR-30a-5p gene proximal promoter region. The miR-30a-5p gene promoter (-2000/+71) was analyzed using the JASPAR database, which gave the relative score. Five c-Myc binding sites at the miR-30a-5p gene promoter are highlighted in red text and gray shadow, and the transcriptional start site is labeled. miR, microRNA.

>rn6\_refGene\_NR\_031843 range=chr9:29541375-29542815 5'pad=0 3'pad=0 strand =+ repeatMasking=none

```
-1260 CGGGGGGTTG GTAGTTGGAG CTGAGTTTGG GATTGACTCA GCTCTGGGCA
-1210
      GCAGTGGTGT ATGAATGCCA AGTGCTAAGC ATGTTTGGAA TGTAGACTTG
                        c-Myc (3.261)
      CCAGCACTGA TAGCGTTTGC TTTGTTCCCC TGTGTGACTT GACCCATTGA
 -1110 GAAATTGTGG TTCCCTAGGA AGTTTTAATA GTTCAAAGTG TTTAGTTAAT
-1060 AAAGAAATG GCCACAACAT AATTCAGCTT GTTTTCACCT GCTGCCTCAA
-1010 AGATTAACTT TCTTAACCAT AATGGGATAG AACTACTTCT TGCTTTGAAA
      AAGAAATTTA TGGAGCTCAT CACATTTGTT AGTTTGCACA AACATGAAGC
      TCTCTGTGGG GATTTTTTAT CATAAAGTCC TAAGGCATTG GCAACTTAAC
 _860
      CTTGTGGTAA CTTCATCTAA AGCCAAATCT CAGGGAAAGG CACACCCACT
        c-Myc (2.312)
 -810 TTTCTGTATC AGCTTAATCT TTCACCAGAC TCAGAGGGTT TTGAATTACT
  -760 TCAGGAAAAG TAGTGACTGA TAGAGCTTAA CCTTCCGTTT TCCTCATTTA
  -710 TTTTCATTAG AGAATGGGAG TGGAATTCAG AACATTCAGA TGTGGAGTTA
                                       c-Myc (2.005)
  -660 ATGTGCGTCA CTGCTGCCAG CCCCTGCTGC CAGCGATAGG TTCTGTGTTT
  -610 TTCCAGTGGG AAACTCTGAT CCACAGACGA TGCATACAGA CCTGCACCTC
  -560 CTTGCTCCTT CTGCTTTTCT CCTGATTGTT ACTTACCCTT TGAAACCTCA
  -510 GCTGACTGTA CAAGGCCTCT GTGGAAGCAG ATTTCAGAAC AAAGGTGATC
  -460 TGTAAAAGTA CCACGTAAGT CTTGTTTCTG TTTCCTGTTC TTATTGATAG
  -410 TTTTAAGTCT CCCTACTTCT GTGTTTAATG TTGAATGTTG AATGTTCTGT
  -360 ACATAATTAA AGCATTTTCC TAGTT<mark>ACATT TGCTC</mark>AAAGA AGGATTTTGA
                                     c-Myc (5.081)
  -310 TGAACTT<mark>ACA CATGTTG</mark>TAG TCCTAGTAAG TCACCTCACC ACTTGCATGA
```

- c-Myc (2.147) c-Mvc (5.019)
- –260 TTGTTTCTTG TGGCCAACAA CAGTAAGCCA TTT<mark>ACATGTG TTG</mark>TTTTCAT c-Myc (10.745)
- -210 AATATGTTTG AGATAGATAA ATAATAATAA ATGTTCCTGA AACATATAAA
- -160 CTATGTCTTA TATAAGCTCT AGTTATATTC TACTAGGGCA TATCTGAACG
- -110 AGGCTTTACA GTTTACAGAA TGTTGCCATT ACATTTTAGA AACACCTACA

Figure S3. Expression of c-Myc and HDAC2 in senescent cardiomyocytes. c-Myc and HDAC2 expression levels were measured via reverse transcription-quantitative PCR and western blotting after cells were transfected with three different Ad-shRNAs of c-Myc or HDAC2 in senescent H9C2 cells. Data are presented as the mean  $\pm$  SD from three independent experiments. \*P<0.05 vs. Ad-shNC group. HDAC2, histone deacetylase 2; Ad-, adenovirus; sh, short hairpin RNA; NC, negative control.

