

Figure S1. CREB expression in LUSC and SCLC tissues. (A and B) CREB expression in (A) LUSC, (B) SCLC and their adjacent normal tissues was measured by ELISA, as presented by scatter plot. Data were analyzed by a paired Student's t-test. CREB, cAMP response element-binding protein; LUSC, lung squamous cell carcinoma; SCLC, small-cell lung cancer.

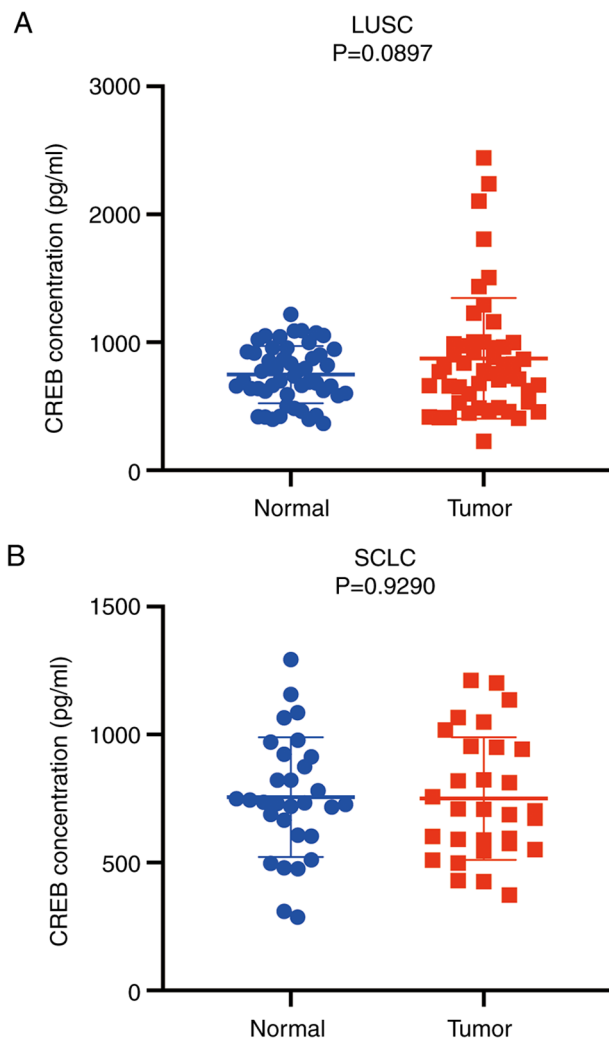


Figure S2. Knockdown efficiency of the EP300-sh1 and -sh2 plasmids. Knockdown efficiency of the EP300-sh1 and -sh2 plasmids compared with their negative empty vector control as measured by immunoblotting in A549 and H1299 cell lines. Representative images from three biological replicates are presented. EP300, E1A binding protein P300; sh, short hairpin

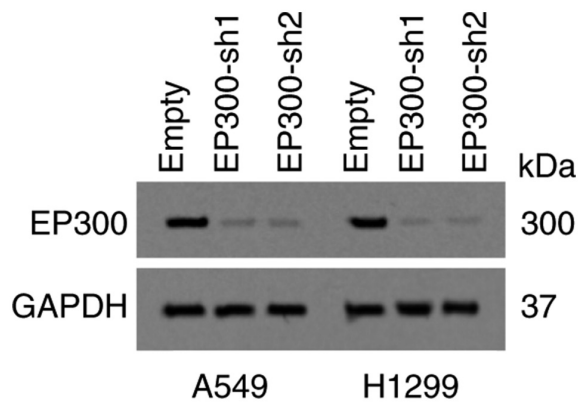


Table SI. Primers used in the study.

## A, Primers used for qPCR

Name	5'-3'
CREB	F: CAGTTATTCAGTCTCCACAAG R: CTTCTCTTCTCAATCCTTGG
GPX4	F: AGTGGATGAAGATCCAACCCAAGG R: GGGCCACACACTTGTGGAGCTAGA
ACSL4	F: GTAATTGGTGGACAGAACATC R: TACTCTCCTGCTTGTAACCTC
CARS	F: GAAGCTCTACAAGGAATATCTG R: GAACCTACATGAACACAACCTC
NRF2	F: CATTCTGAGTTACAGTGTCT R: GGACTACAGTTACCTACTTCTT
HSPB1	F: CAGTCCAACGAGATCACCATC R: ATCCGGGCTAAGGCTTTACTT
SAT1	F: CTTGGTAGCAGAATGGAATGA R: AGAATGGAGGTTGTCATCTAC
EP300	F: ATTAAGGAACTGGAACAGGAG R: AGAGGTCGTTAGATACATTGG
GAPDH	F: ATCATCCCTGCCTCTACTGG R: GTCAGGTCCACCACTGACAC

## B, Primers used for ChIP-qPCR

Name	5'-3'
~-1000 to -751	F: CATTAAAAAGGCAAATCCCTTGGCC R: GTTCAAGCGATTCTCCTGACTCAGCCT
~-750 to -501	F: GTTGCAGTGAGCTGAGATAGC R: GCTCATCCACCATTCTGGCTA
~-500 to -251	F: CTGTTGTCCCAGCTACTCGGGAAGC R: CCTGAGAATACTACTTAAGACTCG
~-250 to -1	F: AGCCGGATAACTGCGCTGCCTC R: GGACGCGCGTCGGCTTTCCGCG
-2k	F: CTTATGCAAGACCAGGATTCGGC R: TCTTAACCTTTTCTGACCCTGCACTG
2k	F: TCCCTGGGGCAGAATGGCTCATG R: ATCCTGTCCCCAGGAGGCCACAAA
CREB	F: AAGCGAGCATGCGCAGTCGCCAA R: GGACGCGCGTCGGCTTTCCGCG

## C, Primers used for promoter analysis

Name	5'-3'
WT-GPX4-Promoter	F: GTACGGTACCATTTACATTA AAAAGGCAAATCCC R: ATTAAGCTAGCGGACGCGCTCGGCTTTCCGCGCCTCCTTT
Mut-GPX4-Promoter	F: GGC GCGAGCGCTCAACA R: TGTTGAGCGCTCGCGCTGGGAGGCCGGAGTGGGAAGAAGGC

## D, Primers used for plasmid construction

Name	5'-3'
CREB-WT-HA	F: ATGCAAGCTTATGACCATGGAATCTGGAGCCGAGAAC R: ATGCGGATCCTTAAGCGTAGTCTGGGACGTCGTATGGGTAAT CTGATTTGTGGCAGTAAAGGTCC
CREB-Del-KID-HA	F: TCAGCACCTGCCATCACTACTGTAA R: GATGGCAGGTGCTGATGTTCCGGAGAAAAGTCTTTTTAAG

Table SI. Continued.

## D, Primers used for plasmid construction

Name	5'-3'
CREB-Del-bZIP-HA	F: ATGCAAGCTTATGACCATGGAATCTGGAGCCGAGAAC R: ATGCGGATCCTTAAGCGTAGTCTGGGACGTCGTATGGGTAT GCTTCTTCAGCAGGCTGTGTAGG
EP300-WT-Myc	F: CTAGAAGCTTATGGCCGAGAATGTGGTGAAC R: CTAGGGTACCCTACAGATCTTCTTCAGAAATAAGTTTTTGTTCG TGTATGTCTAGTGTACTCTGTGAG
EP300-Del-KIX-Myc	F: AGAAAAACGAAGGACCAGACTACAGAAGC R: GGTCTTCGTTTTTCTAGTAGTGGATGGTTGAGCTGCTGTT
EP300-Del-Bromo-Myc	F: AAATACTGCTCCAAGCTCTCTGAGG R: GTAAAGTGCCTCCAAAGTTGGCATCGTAAAGTGCCTCCAAAGT TGGCATC
EP300-Del-CBP/p300-HAT-Myc	F: GACCGCTTTGTCTACACCTGCAATG R: GCAGGTGTAGACAAAGCGGTCATTTTCTTTCCTAGTTCGTGCAC

F, forward; R, reverse; qPCR, quantitative PCR; CREB, cAMP response element-binding protein; GPX4, glutathione peroxidase 4; ACSL4, acyl-CoA synthetase long chain family member 4; CARS, cysteinyl-tRNA synthetase; NRF2, nuclear factor, erythroid 2 like 2; HSPB1, heat shock protein family B small member 1; SAT1, spermidine/spermine N1-acetyltransferase 1; EP300, E1A binding protein P300; ChIP, chromatin immunoprecipitation; WT, wild-type; Mut, mutant.