Figure Supplementary


Figure Supplementary 1. (A-C) Expression of miR-199a-3p as related to tumor size, sex and pathological stage. (D-F) Expression of miR-199a-5p as related to tumor size, sex and pathological stage.


Figure Supplementary 2. A549 and H1299 cells transfected with miR-199a-3p/5p mimic or NC were subjected to wound healing assay and images were taken at 0 h and 24 h .


Figure Supplementary 3. Downregulation of miR-199a-3p/5p could promote the proliferation and migration of NSCLC, and suppress cell apoptosis. (A) Downregulation of miR-199a-3p/5p following transfection with 60 nmol miR-199a-3p/5p inhibitor in A549 and H1299 cells. (B and C) Cell proliferation ability of A549 and H1299 cells transiently transfected with miR-199a-3p/5p inhibitor measured by CCK-8 assay. (D
and E) Colony formation assay in A549 and H1299 cells transfected with miR-199a-3p/5p inhibitor or NC. Representative images and quantitative data are shown. (F and G) A549 and H1299 cells transfected with miR-199a-3p/5p inhibitor or NC were subjected to wound healing assay and images were taken at 0 h and 24 h . (H and I) The rate of apoptosis was analyzed by flow cytometry following transfection with miR-199a-3p/5p inhibitor or NC in A549 and H1299 cells. Each assay was performed in triplicate. $* P<0.05$, $* * P<0.01$, and $* * * P<0.001$.


Figure Supplementary 4. (A and B) There was a negative correlation between miR-199a-3p/5p and Rheb, using GEPIA (http://gepia.cancer-pku.cn/index.html) analysis. (C and D) Rheb expression levels related with tumor stage in LUAD and LUSC tissue samples.


Figure Supplementary 5. (A) Rheb has a positive correlation with mTOR, through

ENCORI (http://starbase.sysu.edu.cn/index.php) database analysis. (B) revealed little change in was decreased In A549 and H1299 cells, the protein expression level of mTOR, p-mTOR after upexpression of miR-199a-3p and miR-199a-5p.

Supplementary Table 1 The clinical-pathological features of 74 cases NSCLC patients

| No. | Gender | Age | Specimen Type | Histologic Type | pTNM |
| :---: | :---: | :---: | :---: | :---: | :---: |
| 1 | M | 47 | Pneumonectomy | Adenocarcinoma | T1bN0M0 |
| 2 | M | 78 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 3 | F | 67 | Pneumonectomy | Adenocarcinoma | T2N1M0 |
| 4 | M | 54 | Lobectomy | Adenocarcinoma | T4N0M0 |
| 5 | M | 49 | Lobectomy | Adenocarcinoma | T2bN2M0 |
| 6 | F | 66 | Pneumonectomy | Adenocarcinoma | T2N2M0 |
| 7 | M | 67 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 8 | F | 62 | Lobectomy | Adenocarcinoma | T1aN0M0 |
| 9 | F | 75 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 10 | F | 58 | Lobectomy | Adenocarcinoma | T2aN1M0 |
| 11 | F | 65 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 12 | M | 72 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 13 | M | 64 | Lobectomy | Adenocarcinoma | T1aN0M0 |
| 14 | F | 62 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 15 | M | 65 | Lobectomy | Adenocarcinoma | T4N2M0 |
| 16 | F | 55 | Lobectomy | Adenocarcinoma | T2aN1M0 |
| 17 | M | 50 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 18 | F | 67 | Pneumonectomy | Adenocarcinoma | T2N1M0 |
| 19 | M | 67 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 20 | M | 71 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 21 | F | 64 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 22 | M | 62 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 23 | M | 58 | Lobectomy | Adenocarcinoma | T2aN0M0 |


| 24 | M | 70 | Lobectomy | Adenocarcinoma | T3N0M0 |
| :---: | :---: | :---: | :---: | :---: | :---: |
| 25 | M | 59 | Lobectomy | Adenocarcinoma | T2aN0M1a |
| 26 | F | 72 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 27 | M | 51 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 28 | M | 51 | Lobectomy | Adenocarcinoma | T1aN0M0 |
| 29 | F | 55 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 30 | F | 53 | Lobectomy | Squamous cell carcinoma | T3N0M0 |
| 31 | M | 54 | Lobectomy | Squamous cell carcinoma | T4N2M0 |
| 32 | M | 58 | Lobectomy | Squamous cell carcinoma | T3N2M0 |
| 33 | M | 65 | Lobectomy | Squamous cell carcinoma | T1bN0M0 |
| 34 | M | 65 | Lobectomy | Squamous cell carcinoma | T2aN2M0 |
| 35 | M | 67 | Lobectomy | Squamous cell carcinoma | T2aN1M0 |
| 36 | M | 67 | Lobectomy | Squamous cell carcinoma | T2aN1M0 |
| 37 | M | 69 | Lobectomy | Squamous cell carcinoma | T2aN0M0 |
| 38 | M | 71 | Lobectomy | Squamous cell carcinoma | T2aN2M0 |
| 39 | M | 61 | Lobectomy | Squamous cell carcinoma | T1bN0M0 |
| 40 | M | 67 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 41 | F | 64 | Lobectomy | Adenocarcinoma | T2aN2M0 |
| 42 | M | 60 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 43 | F | 57 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 44 | M | 65 | Lobectomy | Combined Small Cell | T2bN2M0 |
|  |  |  |  | carcinoma |  |
| 45 | M | 77 | Lobectomy | Squamous cell carcinoma | T2aN0M0 |
| 46 | M | 71 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 47 | F | 64 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 48 | F | 55 | Lobectomy | Adenocarcinoma | T1bN2M0 |
| 49 | M | 61 | Lobectomy | Squamous cell carcinoma | T1bN0M0 |
| 50 | M | 62 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 51 | M | 58 | Lobectomy | Adenocarcinoma | T2aN0M0 |


| 52 | F | 60 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| :---: | :---: | :---: | :---: | :---: | :---: |
| 53 | F | 61 | Lobectomy | Adenocarcinoma | T2aN1M0 |
| 54 | F | 67 | Lobectomy | Adenocarcinoma | T2aN1M0 |
| 55 | F | 57 | Lobectomy | Adenocarcinoma | T4N0M0 |
| 56 | F | 54 | Lobectomy | Adenocarcinoma | T2aN3M0 |
| 57 | F | 66 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 58 | M | 64 | Lobectomy | Adenocarcinoma | T2bN2M0 |
| 59 | M | 49 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 60 | M | 65 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 61 | F | 59 | Lobectomy | Adenocarcinoma | T2aN1M0 |
| 62 | F | 62 | Lobectomy | Adenocarcinoma | T1bN0M0 |
| 63 | F | 45 | Lobectomy | Adenocarcinoma | T2bN2M0 |
| 64 | M | 69 | Lobectomy | Adenocarcinoma | T2bN2M0 |
| 65 | M | 65 | Lobectomy | Adenocarcinoma | T2aN1M0 |
| 66 | F | 58 | Lobectomy | Adenocarcinoma | T2aN1M0 |
| 67 | M | 59 | Lobectomy | Adenocarcinoma | T2N1M0 |
| 68 | M | 60 | Lobectomy | Squamous cell carcinoma | T1bN0M0 |
| 69 | M | 62 | Lobectomy | Adenocarcinoma | T2aN1M0 |
| 70 | F | 55 | Lobectomy | Squamous cell carcinoma | T2aN1M0 |
| 71 | F | 65 | Lobectomy | Squamous cell carcinoma | T2aN2M0 |
| 72 | F | 48 | Lobectomy | Adenocarcinoma | T2aN0M0 |
| 73 | M | 71 | Lobectomy | Adenocarcinoma | T2aN2M0 |
| 74 | F | 55 | Lobectomy | Adenocarcinoma | T2aN0M0 |

Supplementary Table 2: Primer sequences information table

| Name | Sequence ( $5^{\prime}-3{ }^{\prime}$ ) |
| :---: | :---: |
| 18S RNA (F) | CTCGCTTCGGCAGCACA |
| 18S RNA (R) | GCCTCACTAAACCATCCAA |
| U6 snRNA (F) | CTCGCTTCGGCAGCACA |
| U6 snRNA (R) | AACGCTTCACGAATTTGCGT |
| miR-199a-3p qRT | ACAGTAGTCTGCACATTGGTTA |
| miR-199a-5p qRT | CCCAGTGTTCAGACTACCTGTTC |
| Rheb-3'-UTR (F) | GCTCTAGAGGGGAAAGTACAAATACC |
| Rheb-3'-UTR (R) | CGGAATTCATAATCTGAAGGGAGGG |
| 3p-Rheb-3'-mUTR (F) | TGTTACTCAAGTATTAACTAGGCTTCAGTATA |
| 3p-Rheb-3'-mUTR (R) | TTTTACTACGTATAACAATGAGTTCATAATTG |
| 5p-Rheb-3'-mUTR (F) | CTGCTGCAAAGCCTGAGGTCTCAGCGAATA |
| 5p-Rheb-3'-mUTR (R) | GCTTCTTCAGGTAGAATATATTCGCTGAGACCTCAG |
| Rheb qRT (F) | CTATCTTTCCTCAGACATACTCCA |
| Rheb qRT (R) | CACCATATCCAACAATTTGCCATG |
| shRheb-1(F) | CCGGGTTGGTTGGGAATAAGAAAGACTCGAGTCTTTCTTATT |
|  | CCCAACCAACTTTTTG |
| shRheb-1(R) | AATTCAAAAAGTTGGTTGGGAATAAGAAAGACTCGAGTCTT |
|  | TCTTATTCCCAACCAAC |
| shRheb-2(F) | CCGGGCAAGTCTTCATGCTCGGTGACTCGAGTCACCGAGCAT |
|  | GAAGACTTGCTTTTTG |
| shRheb-2(R) | AATTCAAAAAGCAAGTCTTCATGCTCGGTGACTCGAGTCACC |
|  | GAGCATGAAGACTTGC |



Original Figure 3M-Rheb-1


Original Figure 3 M -Rheb-2


Original Figure 3M-Rheb-3


Original figure 3M-GAPDH-1


Original figure 3M-GAPDH-2


Original figure 3M-GAPDH-3


Original Figure 4B-Rheb First


Original Figure 4B-Rheb
Second


Original Figure 4B-Rheb Third


Original Figure 4B-GAPDH-1


Original Figure 4B-GAPDH-2


Original Figure 4B-GAPDH-3


Original Supplementary Figure 5B-p-mTOR First


Original Supplementary Figure 5B-p-mTOR Second


Original Supplementary Figure 5B-mTOR First


Original Supplementary Figure 5B-mTOR
Second


Original supplementary figure 5B-GAPDH-1


Original supplementary figure 5B-GAPDH-2

