Primers	5'3'	Promoter Length (bp)
F1	GCAAGACATTGTGCCAGATGCT	2908 (F1,R1)
F2	CGTGTCACTTCCCAATGTCAGC	2412 (F2,R1)
F3	TGACTGTTGTCCCAAGTGGA	1241 (F3,R1)
R1	TTTCCACGGAGCCTCCGC	

Supplemental Table 1. The primers used to amplify trib2 promoter.

Supplemental Figure 1. miR-206 and MiR-140 induced LTEP-a-2 cell apoptosis

(a) miR-206 inducing cell apoptosis. (b) miR-140 inducing cell apoptosis. Apoptotic cells are shown in the upper left, upper right, and lower right quadrants of each panel. More apoptotic cells were found in the miR-206- or miR-140-treated cells compared with the scrambled oligo control-treated cells after Annexin V-FITC/PI staining. (c) MTT assay. OD was more in miR-206 inhibitor (ASO-206)- (P < 0.01) or miR-140 inhibitor (ASO-140)- treated A549 cells (P < 0.01) compared with miRNA- or scrambled oligo-treated cultures. (d) Colony formation assay. A549 cells, treated with miR-206, miR-140 and controls, were seeded on six-well plates at a density of 500 cells/well. After 12 days, the colonies were fixed with methanol and stained with 0.1% crystal violet (Sigma, St. Louis, MO). ASO-206- or ASO-140 increased colony formation compared with the miRNA or scrambled oligo control treatment. Scrambled, cells treated scrambled control RNA. MiR-206 or miR-140, cells treated with miR-206 or miR-140 oligos. ASO-206 or ASO-140, cells treated with antisense RNA specific to miR-206 or miR-140.



Supplemental Figure 2. Smad3 expression was regulated by miRNAs in LTEP-a-2 cells.

(a) Fluorescence observation. Scale bar = 100 μ M. (b) FACS analysis. MiR-206 and miR-140 treatment reduced the number of GFP positive cells and intensity of GFP fluorescence significantly compared with control oligos treatment. (c) Smad3 and p-Smad3 protein (55kD) expression were detected by western blot. The results showed that p-Smad3 and Smad3 expression was inhibited by miR-140 or miR-206 treatment compared with control treatment. The Smad3/actin (or p-Smad3/actin) was shown on the right of the gel. ** p<0.01. miRNA treatment *vs*. control treatment. scrambled, cells treated scrambled control RNA. MiR-206 or miR-140, cells treated with miR-206 or miR-140 oligos. Mu-206 or Mu-140, cells treated with mutation sequence of miR-206 or miR-140. SiRNA-smad3, small interfering RNA specific to knock down Smad3 expression.



Supplemental Figure 3. The effect of TGF-\$1 on miR-140 or miR-206 expression in

LTEP-a-2 cells. (a) miR-140, (b) miR-206. The levels of miR-140 (A) and miR-206 were obviously decreased in TGF- β 1-treated compared with untreated LTEP-a-2 cells. **p<0.01. n = 3 replicates.



Supplemental Figure 4. TRIB2 expression in LTEP-a-2. TRIB2 expression (46kD) was decreased obviously in miR-140- or miR-206-treated cells. Relative values for TRIB2 vs Actin are indicated to the right of the gel. **p<0.01. *vs.* negative control. n = 3 replicates. MiR-206 or miR-140, cells treated with miR-206 or miR-140 oligos. Mu-206 or Mu-140, cells treated with mutation sequence of miR-206 or miR-140. SiRNA-Smad3, cells treated with small interfering RNA specific to Smad3. The sense sequence of siRNA-Smad3,

5'-CACAUAAUAACUUGGACCUU -3'.



Supplemental Figure 5. Mouse TRIB2 promoter activity analysis. TGF-β1 treatment increased mouse trib2 promoter activity, but SB431542 blocked TGF-β1-mediated mouse trib2 promoter activity. **p<0.01 vs. TGF-β1 treatment.



Mouse trib2 promoter

Supplemental Figure 6. Mutant of "CAGACA" box. The "CAGACA" box was missed in mutation promoter compared with wide type promoter.



Supplemental Figure 7. The expression of E-cadherin and α -SMA in LTEP-a-2 cells. Western Blot showed that the expression of E-cadherin (130kD) was increased, while α -SMA (45kD) level was reduced in miR-140- (or miR-206-) treated LTEP-a-2 cells. Relative values for E-cadherin or α -SMA vs Actin are indicated to the right of the gel. **p<0.01. Scrambled, cells treated scrambled control RNA. MiR-206 or miR-140, cells treated with miR-206 or miR-140 oligos. SiRNA-Smad3 or SiRNA-control, cells treated with small interfering RNA specific to Smad3 or siRNA-control. TGF- β 1, 10ng/ml TGF- β 1 treatment. SB431542, 5 µmol/ml SB431542 treatment.

