

1 **Supplementary Figure legends**

2 **Supplementary Figure 1. A.** The mRNA expression level of miR-21 in human colorectal cancer was  
3 analyzed using the Oncomine TCGA Colorectal dataset. The mRNA expression level of VMP1 in human  
4 colorectal cancer were analyzed using the Oncomine Skrzypczak Colorectal dataset. **B.** MTT assay to  
5 calculate the concentration of 5-FU resistance achieved in case of HCT116-FUR cells. Data are presented  
6 as mean $\pm$ SD. \*\*\* $P$ <0.001.

7 **Supplementary Figure 2. A.** The primer binding sites of VMP1 and miR-21 transcripts. RT-PCR primers  
8 are depicted as facing arrows, reverse transcription primer is depicted as single thick arrow. **B.** HCT116  
9 was starved with glucose-free medium for 2h and 4h, the expression of miR-21, and the amount of pri-miR-  
10 21 and VMP1-miR-21 transcripts, were measured by qPCR. The protein expression level of VMP1 was  
11 measured by Western blot. **C.** HCT116 was treated with 5 $\mu$ M rapamycin for 12h, 24h and 48h, the  
12 expression of miR-21, and the amount of the pri-miR-21 and VMP1-miR-21 transcripts, were measured by  
13 qPCR. The protein expression level of VMP1 was measured by Western blot. Data are presented as mean  
14  $\pm$ SD. \*\*\* $P$ <0.001, \*\* $P$ <0.01, \* $P$ <0.05.

15 **Supplementary Figure 3. A.** RKO and HCT116 were transfected with miR-21 nucleotide mimic, and cells  
16 were treated with 1 $\mu$ M Afuresertib after transfection for 24h, the protein level of TFEB in cytoplasm and  
17 nucleus respectively was measured by Western blot, H3 and  $\beta$ -actin were used as nuclear and cytosolic  
18 markers, respectively. **B.** The protein expression level of TFEB and VMP1 were measured by Western blot  
19 after over-expression of the constitutively active TFEB by pCIP-caTfeb vector or knock-down of TFEB

20 expression by siRNA in HCT116 cells. **C.** The predicted binding sites of AP-1 and AR on VMP1 promoter  
21 region were shown.