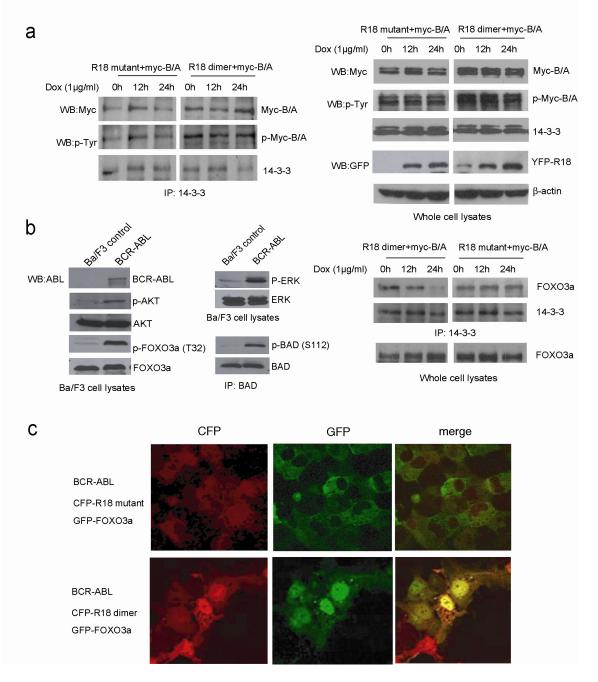
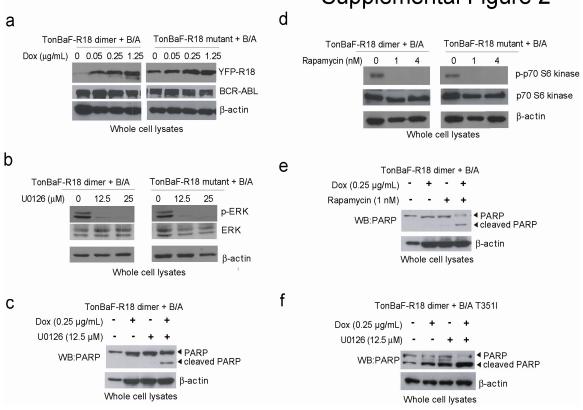
Supplemental Figure 1. R18 interferes with interaction between 14-3-3 and FOXO3a, but not 14-3-3/BCR-ABL association or BCR-ABL kinase activation in Ba/F3 cells. (a) Myc-tagged BCR-ABL transformed TonBaF cells inducibly expressing YFP-tagged R18 dimer or mutant were cultured with Dox in the absence of IL-3. Left: Coimmunoprecipitated Myc-BCR-ABL in 14-3-3 immunoprecipitates was detected by Western blotting. *Right*: Phosphorylation levels of Myc-BCR-ABL and expression levels of BCR-ABL, 14-3-3 and YFP-R18 in the cell lysates were probed by Western blotting as controls. (b) R18 dimer dissociates 14-3-3/FOXO3a association. Left: Expression of BCR-ABL activates both AKT and ERK pathways that results in phosphorylation of FOXO3a and BAD, respectively, which are consequently sequestered by 14-3-3 binding. *Right*: BCR-ABL transformed TonBaF cells inducibly expressing YFP-tagged R18 dimer or mutant were cultured with Dox in the absence of IL-3. Co-immunoprecipitated FOXO3a in 14-3-3 immunoprecipitates were detected. (c) Expression of R18 dimer inhibits cytoplasmic sequestration of FOXO3a by BCR-ABL, and rescues nuclear localization of FOXO3a. COS7 cells were co-transfected with BCR-ABL, CFP-tagged R18 dimer or mutant, and EGFP-FOXO3a. The molar ratio of DNA amount in the cotransfection experiment was <u>BCR-ABL:R18:FOXO3a = 2:1:1</u> to ensure successful BCR-ABL transfection in cells that are both CFP and EGFP positive.

Supplemental Figure 2. (a) Induction of R18 expression by Dox was detected by Western blotting. (b) Inhibition of ERK phosphorylation by U0126 treatment in TonBaF-R18 cells expressing BCR-ABL. (c) Combination of U0126 treatment and R18 expression enhanced cleavage of PARP in BCR-ABL-transformed TonBaF-R18 cells. (d)

Inhibition of mTOR activity as assessed by phosphorylation level of p70 S6 kinase by rapamycin. (e) Combination of rapamycin treatment and R18 expression enhanced cleavage of PARP in TonBaF-R18 BCR-ABL cells. (f) Treatment with U0126 along with R18 expression enhanced cleavage of PARP in BCR-ABL T315I mutant-transformed TonBaF-R18 cells.

Supplemental Figure 1





Supplemental Figure 2