

Data S1. To determine the dose of 3-methyladenine (3-MA) for inhibiting autophagy without obvious cytotoxicity in CRC cells, RKO cells were exposed to 3-MA alone or combined with 40 μ M PRIMA-1^{met} for 48 h followed by a Cell Counting Kit-8 assay. The results showed that cell proliferation decreased after the treatment of 3-MA in a dose-dependent manner once over 800 μ M (Fig. S1). However, the western blotting results indicated that dosages lower than 800 μ M (200 and 400 μ M) for 3-MA had no effect on the expression

of LC3-II (The figures are not shown in the manuscript). To further make the working dose of 3-MA more reliable, a colony formation assay was performed in the HCT116^{wt} cell line. Cell colonies were observed and counted following 10 days of treatment with PBS control or 3-MA at indicated dosages of 800 and 3000 μ M. As predicted, the number of colonies was significantly suppressed with 3-MA at the high dose of 3000 μ M compared to that observed in cells treated with PBS alone ($P < 0.01$; Fig. S2).

Figure S1. RKO cells were treated with 40 μ M PRI and 3-MA at the indicated dosages alone or in combination for 48 h, followed by a Cell Counting Kit-8 assay. The percentage of cell proliferation was normalized with the respective PBS control. 3-MA alone suppressed cell proliferation once the dose was over 800 μ M. Treatment administered in combination resulted in a significant suppression to the effect of PRIMA-1^{met} with the dosages of 3-MA from 100 to 800 μ M. Data were obtained from triplicate samples and assessed as the mean + SEM. 3-MA, 3-methyladenine; PRI, p53-reactivation and induction of massive apoptosis-1, APR-017 methylated.

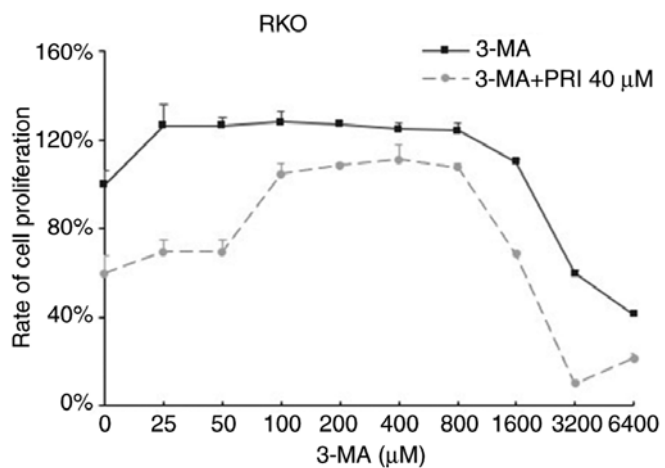


Figure S2. HCT116^{wt} were seeded into 6-well plates in duplicate following treatment with PBS (ctrl) or 3-MA at indicated dosages of 800 and 3,000 μM for 48 h. After 10 days, cell colonies were observed, imaged and analyzed using ImageJ version 1.8.0 software (National Institutes of Health). Data are presented as the mean + SEM (n=2). **P<0.01, compared with the PBS-treated group (ctrl). 3-MA, 3-methyladenine.

