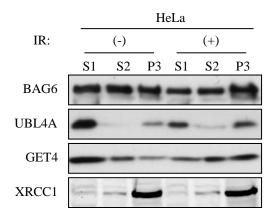
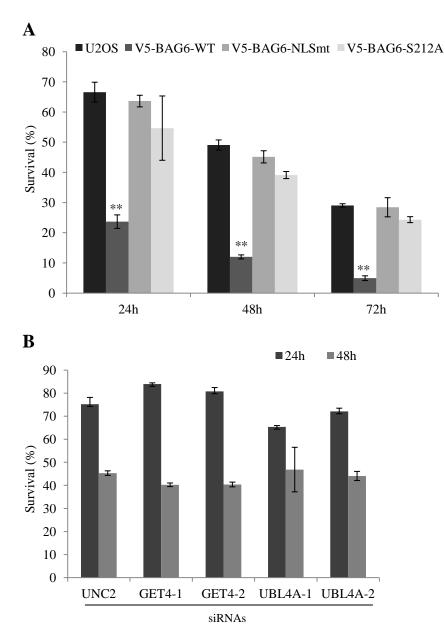


- – no inhibitor
- C- chloroquine (lysosomal inhibitor), 200  $\mu M$
- L-lactacystin (proteosomal inhibitor), 20  $\mu M$
- N-NH4Cl (lysosomal inhibitor), 20 mM

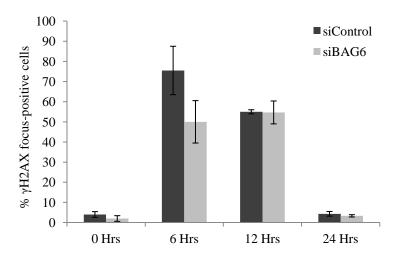
Supplemental Figure 1. **GET4 and UBL4A can not be stabilized upon proteosome and lysosome treatment in the absence of BAG6**. U2OS (A) and HeLa (B) were treated with indicated inhibitors at indicated final concentrations for 6 hrs. Cells were harvested and protein distributions were assessed by Western blotting using indicated antibodies .



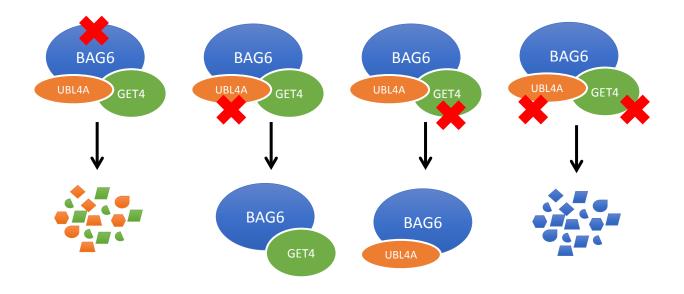
Supplemental Figure 2. **UBL4A and GET4 translocate to nucleus upon DNA damage.** Chromatin fractionations were performed on untreated or IR-treated HeLa cells. Protein distributions were assessed by Western blotting using indicated antibodies. S1 fraction – cytoplasm, S2 – soluble nucleus fraction and P3 – chromatin bound fraction.



Supplemental Figure 3. Cells are not resistant to DNA damage induced cell death in the absence of GET4 or UBL4A. A, U2OS cells and U2OS cells stably expressing the indicated proteins were treated with doxorubicin (2  $\mu$ g/ml). Cell survival was determined using WST-1 assays at 24, 48 or 72 hours post treatment. B, U2OS cells transfected with indicated siRNAs were treated with doxorubicin (2  $\mu$ g/ml). Cell survival was determined as described in (A). All error bars represent mean ± SD (n=3). \*\* - p<0.01.



Supplemental Figure 4. **BAG6 complex does not affect DNA damage-induced**  $\gamma$ **H2AX foci formation in U2OS cells depleted of BAG6 complex.** Quantitative determination of cells forming  $\gamma$ H2AX foci upon DNA damage. U2OS cells were transfected with BAG6 siRNA and treated with 10 Gy IR. Cells were fixed at indicated times after IR and stained with  $\gamma$ H2AX antibody and subjected to quantification. At least 300 cells were counted to determine the percentage of foci-containing cells. Data are represented as mean ± SD (n=3).



Supplemental Figure 5. **BAG6 complex stability.** BAG6 complex exists as either a stable ternary complex composed of BAG6, UBL4A and GET4, or binary complex composed of BAG6/UBL4A or BAG6/GET4. In the absence of BAG6, GET4 and UBL4A levels are reduced; in the absence of both GET4 and UBL4A, BAG6 level is reduced. The mechanism that regulate the protein levels is yet to be identified.