

**Supplemental Figure 1**: Phase contrast images of MDA-MB-231 and HMEC cells treated with indicated concentrations of ManA for 9h. A) Dose dependent induction of cytoplasmic vacuolation by ManA in MDA-MB-231cells. B) Man A failed to induce cytoplasmic vacuolation in HMEC.



**Supplemental Figure 2:** Failed inhibition of cell death by autophagy inhibitor chloroquine. MTT assay with ManA (5uM), Chloroquine (50uM, 100uM) individually and together in BT-20 (A) and HCC1937 (B) cells at 24h, 48h, 72h. C) Western blots for Bip, p62, CHOP, LC3 and protein ubiquitination in total cell lysates of MDA-MB-231 cells treated with ManA without or with Chloroquine.



**Supplemental Figure 3**: Western blot of total cell lysates from ManA treated MDA-MB-231 cells and Saturosprin (STA) treated Hela cells with Caspase-3 antibody that recognizes cleaved Caspase3. No cleavage of Caspase-3 was seen in ManA treated MDA-MB-231 cells.



**Supplemental Figure 4**: Confocal images of control (Con) and ManA treated MDA-MB-231 cells using LC3 antibody (red). Nuclei were labeled with DAPI (blue).



Supplemental Figure 5: A) ManA induced cytoplasmic vacuolation in MDA-MB-231 cells is prevented by general transcription and translation inhibitors Actinomycin D (ActD, 8 µM) and cycloheximide (CHX, 25 µM) respectively. B) Western blots of MDA-MB-231 total cell lysates with ManA treatment for Bip, p62, LC3 and ubiquitinated proteins in the absence and presence of ActD and CHX.



MDA-231\_ MDA-231\_ Control shRNA LC3 shRNA



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**Supplemental Figure 6:** LC3 knockdown reduced colony formation by MDA-MB-231 cells in soft Agar. A) Phase contrast image control shRNA and LC3shRNA expressing MDA-MB-231 cells. B) Crystal violet stained colonies of control shRNA and LC3 shRNA expressing MDA-MB-231 cells in soft Agar. C) Graphic representation of relative number of colonies formed by MDA-MB-231 cells expressing control shRNA and LC3 shRNA and LC3 shRNA (\* p<0.001).



**Supplemental Figure 7**: LC3 deficiency inhibited migration properties of MDA-MB-231 cells. A) Phase contrast images of toluidine blue stained transit well migration assay of con shRNA and LC3 shRNA MDA-MB-231 cells. B) Comparative differences in migration index of MDA-MB-231 cells expressing con shRNA and LC3 shRNA (\* p<0.001).



Supplemental Figure 8: LC3 deficiency inhibited invasion properties of MDA-MB-231 cells.

A) Toluidine blue stained phase contrast images of MDA-MB-231 cells expressing con shRNA and LC3 shRNA invading through Matrigel coated insert membranes. B) Comparative differences in invasion index of MDA-MB-231 cells expressing con shRNA and LC3 shRNA (\*p<0.001).