

Supplementary Figure-1- A- Immunofluoresence photomicrograph of F-Actin staining (green) of ZR-75-1, MCF-7 cells and their radioresistant counterparts. Scale bar- 10μm. **B**- Bar graph showing the platting efficiency. C- Bar graph showing the quantification of γH2AX level in MCF-7 and MCF-7^{FR} cells. D- Immunofluoresence photomicrograph of γH2AX foci in ZR-75-1 cells and ZR-75-1^{FR} cells at different time points post 2Gy irradiation. Red- γH2AX foci and Blue-DAPI (Scale bar-5μm). E- Line graph represents the quantification of γH2AX level from panel D. Statistical significance between the two groups was determine by Anova.



Supplementary Figure-2-A- Table showing the level of CSC sub-population identified by side population assay. B- Dot plot showing the level of ALDH positive cells in ZR-75-1, MCF-7 cells and their radioresistant counterpart. Inset dot plot shows the level of ALDH positive cells in presence of DAEB. C- Graph showing the number of spheroids formed by SP, MP and NSP of ZR-75-1 and MCF-7 cells. D- Immunofluoresence photomicrograph of γ H2AX foci in MP, SP and NSP of ZR-75-1 and MCF-7, 1 hour post 2Gy irradiation, where red indicates γ H2AX foci, and blue indicates DAPI (Scale bar-10µm). E- Bar graph represents the quantification of γ H2AX

level from panel D. F, G- Bar graph showing the platting efficiency of ZR-75-1 and MCF-7 cells. Statistical significance of data in C and E is determined by Student's t-test. **- P<0.01, ***-P<0.001, ****- P<0.0001



Figure 3- A- Graph representing the relative level of EpCAM mRNA to that of housekeeping gene GAPDH in CSC (SP) and non-CSC (NSP) of ZR-75-1 and MCF-7. **B**, **C**-Immunofluoresence photomicrograph and graphical representation of the level of EpCAM(red) in CSC and non-CSC of ZR-75-1 and MCF-7 where nucleus is represented in blue. Scale bar -5µm. Statistical significance of data in A and C is determined by Student's ttest. *- P<0.05, ****- P<0.0001



Supplementary Jigure 4-A-Dot plot showing the distribution of MCF-7 cetts based on the EpCAM detected using flowcytometry. The dot plots in red show the secondary control, while one in blue show the positive control. Cells isolated as $EpCAM^{Low}$ and $EpCAM^{High}$ is shown in black box. **B**,**C**- Immunofluoresence photomicrograph and graphical representation of the level of EpCAM(red) in $EpCAM^{Low}$ and $EpCAM^{High}$ population of MCF-7 cells where nucleus is represented in blue. Scale bar $-10\mu m$. **D** -Semi-quantitative western blot showing expression level of EpCAM in $EpCAM^{Low}$ and $EpCAM^{High}$ population of MCF-7 cells. **E**- Comparison of survival fraction between $EpCAM^{Low}$ and $EpCAM^{High}$ population of MCF-7 cells against different doses of radiation. Inset table indicates radio-resistivity index (D₀ value) for each. The dose modifying factor(DMF) is also mentioned in inset. Statistical significance of data in C, E and F is determined by Student's t-test.*- P<0.05, **- P<0.01, ****- P<0.001



Supplementary Figure-5-A- Bar graph showing the platting efficiency of ZR-75-1, ZR-75-1^{FR} and ZR-75-1^{EpCAM} cells. **B**- -Immunofluoresence photomicrographs of γ H2AX foci in ZR-75-1 and ZR-75-1^{EpCAM} cells after treatment with doxorubicin, cisplatin and gemcitabine for 24hours. Scale bar is 10µm. **C** -**F** Line graph showing the growth curve without irradiation and on 2Gy, 4Gy and 8Gy irradiation, respectively. **G**- Cell cycle analysis of ZR-75-1, ZR-75-1^{FR} and ZR-75-1^{EpCAM} cells at different time points, post 10Gy irradiation. Statistical significance of data in C-F is determined by Anova. *- P<0.05, **- P<0.01, ***- P<0.001, ****- P<0.001



Supplementary Figure-6-A- Diagram showing the injection of cells. 100,000 ZR-75-1 and ZR- $75-1^{EpCAM}$ cells were injected in the mammary fatpad. **B**- Table showing the number of mice in which there is palpable tumor after 30days. **C**- Graph representing the volume of tumor (from panel **B**).



Supplementary Figure-7- Representative photomicrograph image of lungs of mice belonging to ZR-75-1 tumor group (A) and ZR-75- 1^{EpCAM} tumor group (B). The images are in 1X magnification. A part of 1X, which is marked with blue box, is further imaged at 5X

magnification. A part of 5X image marked with black box was also further imaged at 10X magnification. The black arrows denote the secondary metastasis within the lung.