

Figure S9

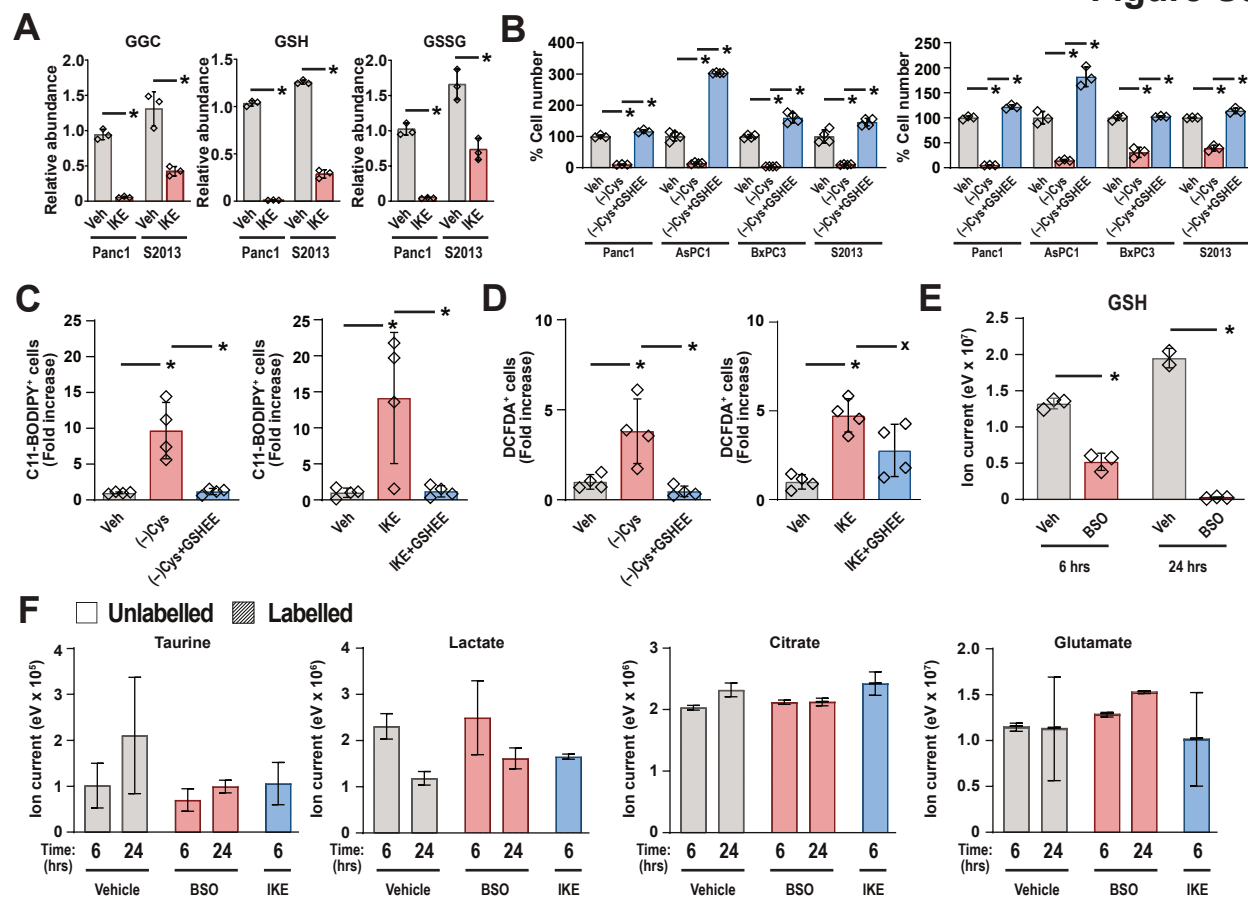


Fig. S9. Metabolomics analysis of cysteine utilization and rescue of cell death by cell permeable GSH. (A) Measurement of gamma glutamyl cysteine (gGC), GSH, and GSSG in two sensitive cell lines treated with 0.1% DMSO (Veh) or 5 μ M IKE for six hours. Data are mean \pm SD with $n = 3$ biological replicates. * $p < 0.05$, Student's t-test. (B) GSH-EE rescue of cystine withdrawal and system x_c^- inhibition in sensitive cell lines. Data are mean \pm SEM with $n = 3$ independent experiments. * $p < 0.05$ by Tukey's test. (C and D) Flow cytometric analyses of sensitive cell lines grown without cystine (-cys) or with 5 μ M IKE and in the presence of GSH-EE for 6-8 hours. Cells were stained with C-11 BODIPY and DCFDA to mark lipid oxidative stress and general oxidative stress, respectively. Data are means \pm SD with $n = 4$ independent cell lines tested in triplicate. * $p < 0.05$ by Tukey's test. x = not significant. (E) GSH levels as measured by mass spectrometry in PANC-1 cells treated with vehicle or 600 μ M BSO for listed timepoints. Data are means \pm SD with $n =$ at least 2 biological replicates. * $p < 0.05$ by Student's t test. (F) Measurement of labeled and unlabeled levels of taurine, lactate, glutamate, and citrate in PANC-1 treated with vehicle (0.05% DMSO), 5 μ M IKE, or 600 μ M BSO for indicated times. (All comparisons not significant by Tukey's test). No labeled species for these metabolites were detected.