

## Supplementary Figure S2.

Cystathionine and homocysteine rescue NSCLC cell viability under cystine starvation (related to Fig. 1). **A,** Measurement of relative NSCLC cell number under cystine-replete (200  $\mu$ M) or cystine-starved (0  $\mu$ M) conditions supplemented with 1 mM cystathionine or 1 mM homocysteine. **B,** Measurement of relative NSCLC cell number under cystine-starved condition treated with cystathionine or homocysteine in the presence of absence of 100  $\mu$ M of propargylglycine. Relative cell number was determined by crystal violet staining, followed by normalization to cystine-replete, untreated conditions. For **A** and **B**, data are presented as mean  $\pm$  SD. **C,** Labeled fraction of serine and concentration of homocysteine, cystathionine, cysteine, and glutathione after 24 h  $^{13}$ C<sub>3</sub>-serine labeling in H2009 under cystine-replete (200  $\mu$ M) or cystine-starved (0  $\mu$ M) condition supplemented with 1 mM cystathionine or 1 mM homocysteine. N=3 biological replicates for each condition. \*p < 0.05, \*\*p < 0.01, \*\*\*p < 0.001, and \*\*\*\*p < 0.0001. Cys, cysteine; (Cys)<sub>2</sub>, cystine; Cth, cystathionine; Hcy, homocysteine; PPG, propargylglycine; GSH, glutathione; Ser, serine.