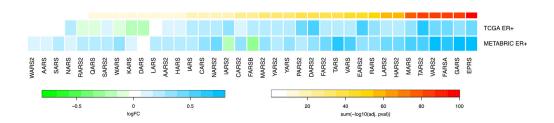
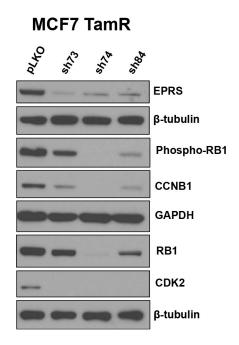
EPRS is a critical regulator of cell proliferation and estrogen signaling in ER+ breast cancer

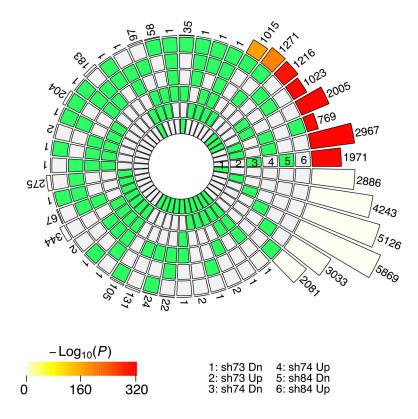
Supplementary Materials



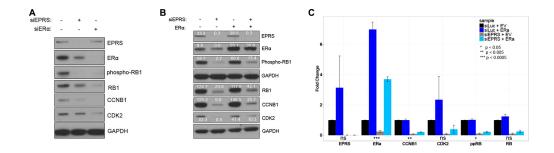
Supplementary Figure S1: Aminoacyl tRNA synthetase expression in ER+ breast cancer. Differential expression of ARS genes in ER+ breast tumors compared to adjacent normal samples in TCGA and METABRIC cohorts. Genes are ordered by the sum of their –log10(adjusted *p* values) from each individual dataset.



Supplementary Figure S2: *EPRS* **shRNA downregulates cell cycle proteins in MCF7 TamR cells.** Representative western blot of shEPRS-induced downregulation of cell cycle proteins in MCF7 TamR cells.



Supplementary Figure S3: shEPRS RNA-seq. *EPRS* shRNA RNA-sequencing differentially-expressed genes (FDR < 0.05, FC > 1.3). Overlaps between shRNAs are tested and visualized using the SuperExactTest.



Supplementary Figure S4: *EPRS* **affects proliferation partially through** *ESR1*. (A) *ESR1* knockdown phenocopies *EPRS* knockdown. Representative immunoblot shown. (B and C) Rescue of *ESR1* expression partially rescues siEPRS-induced downregulation of cell cycle proteins. Error bars represent SEM. siLuc: luciferase-targeting siRNA (control). siEPRS: pool of four unique siEPRS-targeting siRNAs.

 $\label{thm:supplementary} \textbf{Supplementary Table S1: Tamoxifen response and nonresponse genes.}$

See Supplementary_Table_S1

Supplementary Table S2: EES keydrivers. "1" indicates affirmative, "0" indicates negative. "Survival.cox" and "survival.km" indicate Cox regression and Kaplan-Meier, respectively, p-values combined from TCGA, METABRIC, Miller and Wang datasets using Fisher's method. See Supplementary_Table_S2