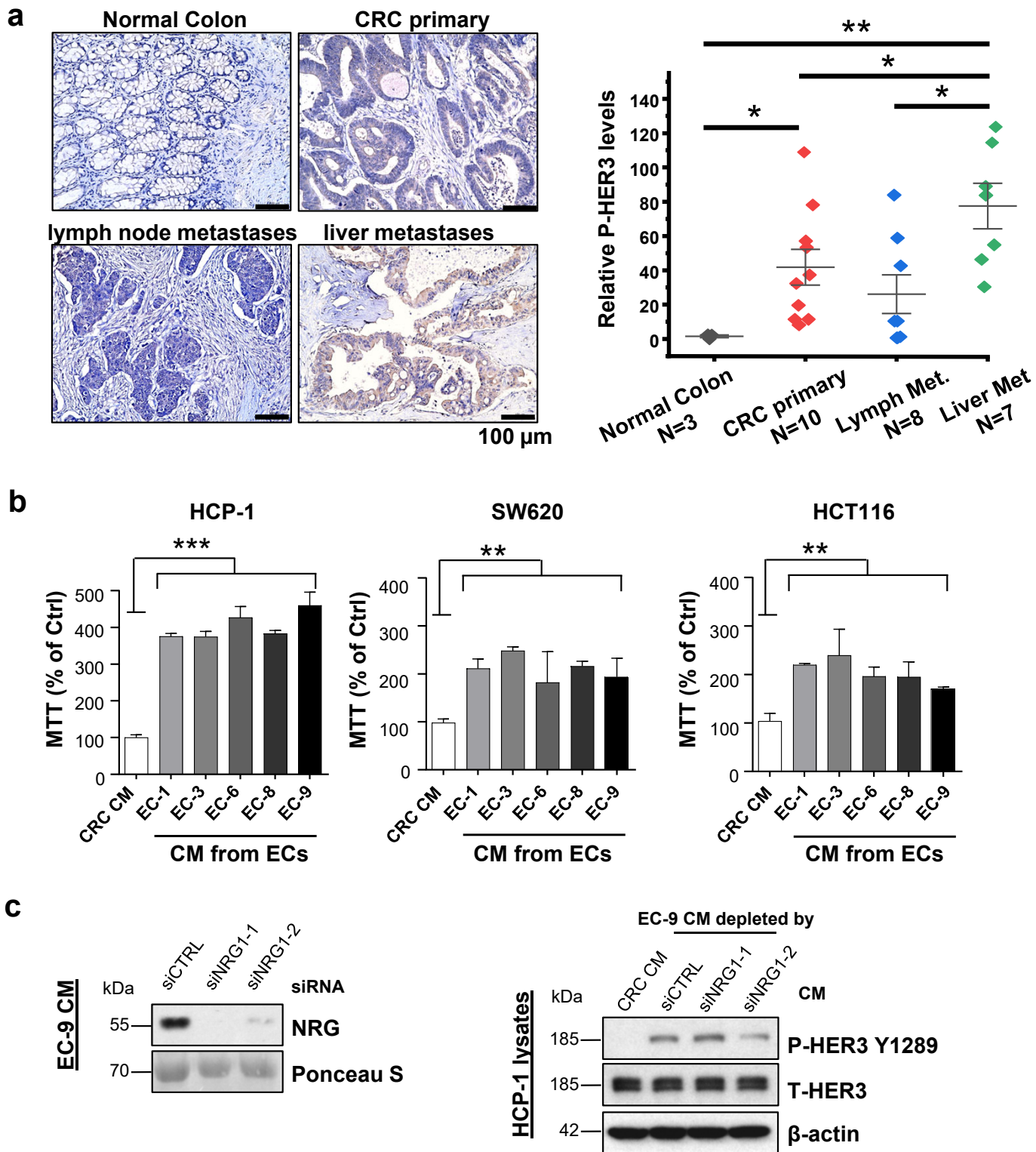


# Supplementary Figure 1



**Supplementary Figure 1. HER3 is activated in mCRC independent of canonical ligand neuregulins.** (a) Representative images of IHC staining and quantifications showing phospho-HER3 (P-HER3, brown) at Y1289, counterstained with H&E, in human microarray colon mucosa and CRC tissues from different stages. Sections were counter stained with hematoxylin. Scale bars 100  $\mu$ m. (b) The MTT assay showing relative cell viability in CRC cells treated by conditioned medium (CM) from each CRC cell line or from different EC lines. (c) Western blots showing levels of neuregulins (NRG) in CM from EC-9 cells with control (siCTRL) or neuregulin-specific siRNAs (siNRG1-1 and -2. Other NRG isoforms not expressed in ECs), and P-HER3 levels in CRC cells treated by CRC CM, or CM from EC-9 with indicated siRNAs. Data in (A) are shown as Mean  $\pm$  SD, \* $p$ <0.05, \*\* $p$ <0.005 by one-way ANOVA test, in (B) are shown as Mean  $\pm$  SEM, \*\* $p$ <0.005, \*\*\* $p$ <0.0005 by unpaired two-tailed  $t$ -test. Ponceau S, total HER3 (T-HER3), and  $\beta$ -actin are shown as loading controls.