SUPPLEMENTARY INFORMATION

“ADAM8 expression in invasive breast cancer promotes tumor dissemination and metastasis” by Romagnoli et al.

Supplementary Figures:

Supplementary Fig S1. ADAM8 mRNA levels correlate with clinicopathological variables of human breast tumors.

Supplementary Fig S2. ADAM8 proform is induced in Hs578T cells grown in 3D-cultures, which co-migrates with the one seen in MDA-MB-231 cells.

Supplementary Fig S3. Ectopic ADAM8 expression rescues the invasive phenotype of stable ADAM8 knockdown cells.

Supplementary Fig S4. ADAM8 knockdown tumors fail to grow beyond a palpable size and are poorly vascularized in a mammary fat pad mouse model.

Supplementary Fig S5. HIF-1α is induced in breast cancer cell lines under hypoxia.

Supplementary Fig S6. Ectopic expression of full-length ADAM8 and remnant form in stable ADAM8 knockdown cells.

Supplementary Fig S7. siRNA-mediated knockdown of ADAM8 in MDA-MB-231 cells reduces their ability to adhere to and transmigrate through HUVECs.

Supplementary Fig S8. ADAM8 is not detected in HUVECs.

Supplementary Fig S9. Anti-ADAM8 antibody inhibits the metalloprotease activity of ADAM8.

Supplementary Fig S10. VEGF-A mRNA levels are unchanged in stable ADAM8 knockdown cells.
**Supplementary Tables:**

**Supplementary Table S1.** Multivariate analysis of *ADAM8* mRNA as a predictor in breast cancer.

**Supplementary Table S2.** Clinical characteristics of the human primary breast tumors analyzed by ADAM8 ELISA.

**Supplementary Table S3.** Clinical characteristics of the sera from breast cancer patients analyzed by ADAM8 ELISA.

**Supplementary Table S4.** Clinical characteristics of the primary and metastatic samples from breast cancer patients analyzed for ADAM8 by immunohistochemistry.

**Supplementary Methods:**

**Cell Lines and Culture Conditions**

Western blotting

ATP assay

Soft agar and Matrigel outgrowth assays

Migration/invasion and transendothelial migration assays

Tube formation assay

ADAM8 metalloproteinase activity assay

Detection of CTCs

**References**