



Supplementary Materials

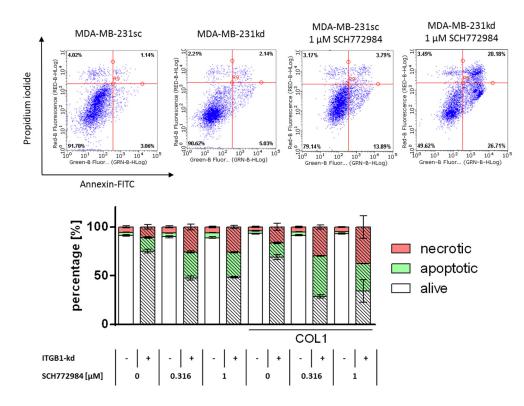


Figure S1. Annexin V-FITC/Propidium iodide assay of SCH772984 (0.316 μ M and 1 μ M, for 48 h) incubated MDA-MB-231 scrambled (sc) and integrin β₁-knockdown (ITGB1-kd) cells in the presence and absence of collagen type 1 (COL1, n = at least 3 biological samples).

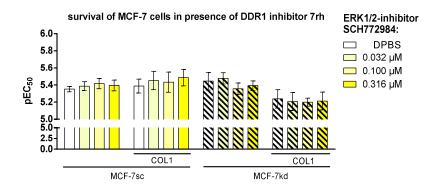


Figure S2. Combination effect of 7rh and SCH772984 cytotoxicity in MCF-7 scrambled (sc) and integrin $β_1$ -knockdown (kd) cells in the presence and absence of collagen type 1 (COL1), displayed by the pEC₅₀ values of 7rh cytotoxicity in dependence of SCH772984 (n = at least 3 biological samples). MCF-7 cells showed no sensitization upon this combination treatment. Statistical analysis was performed via unpaired t-test.

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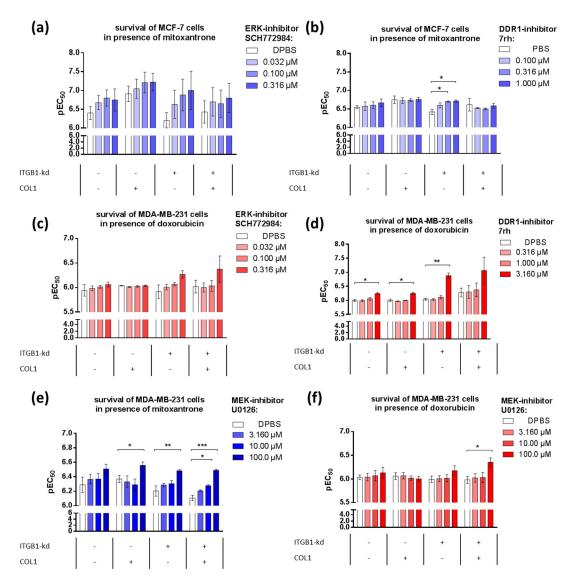


Figure S3. The impact of ERK1/2, MEK1/2 and DDR1-inhibitors on the cytotoxic treatment of MDA-MB-231 and MCF-7 scrambled and integrin $β_1$ -knockdown (ITGB1-kd) cells with doxorubicin or mitoxantrone in combinatory approaches in the presence and absence of collagen type 1 (COL1). (a,b) The ERK1/2-inhibitor SCH772984 (a) and DDR1-inhibitor 7rh (b) were combined with mitoxantrone in MCF-7 cells. (c,d) The ERK1/2-inhibitor SCH772984 (c) and DDR1-inhibitor 7rh (d) were combined with doxorubicin in MDA-MB-231 cells. (e,f) The MEK1/2-inhibitor U0126 was combined with (e) mitoxantrone and (f) doxorubicin in MDA-MB-231 cells. The cytostatics were applied in increasing concentrations to gain a pEC50 value. The inhibitors were additionally added at a constant concentration. Shown are means of pEC50 values and statistical analysis was performed using unpaired t-test (*p < 0.05; **p < 0.01, p = 3 biological samples).

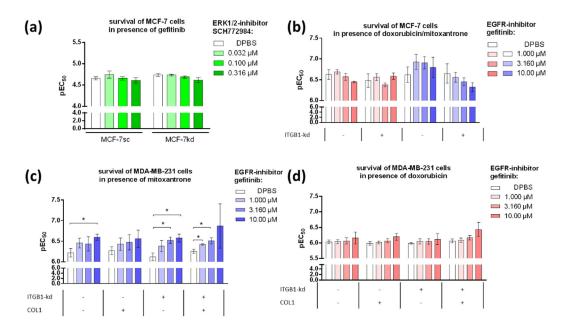


Figure S4. (a) Survival of MCF-7 scrambled (sc) and integrin $β_1$ -knockdown (ITGB1-kd) cells in the presence of gefitinib in dependence of SCH772984. Inhibition of EGFR by gefitinib had no further synergistic effect with ERK1/2-inhibition in MCF-7 cells. (b) Survival of MCF-7 and (c,d) MDA-MB-231 sc and kd cells in the presence and absence of collagen type 1 (COL1). The cells were incubated with mitoxantrone or doxorubicin in dependence of EGFR inhibitor gefitinib. Statistical analysis was performed via unpaired t-test (n =at least 3 biological samples, *p < 0.05).

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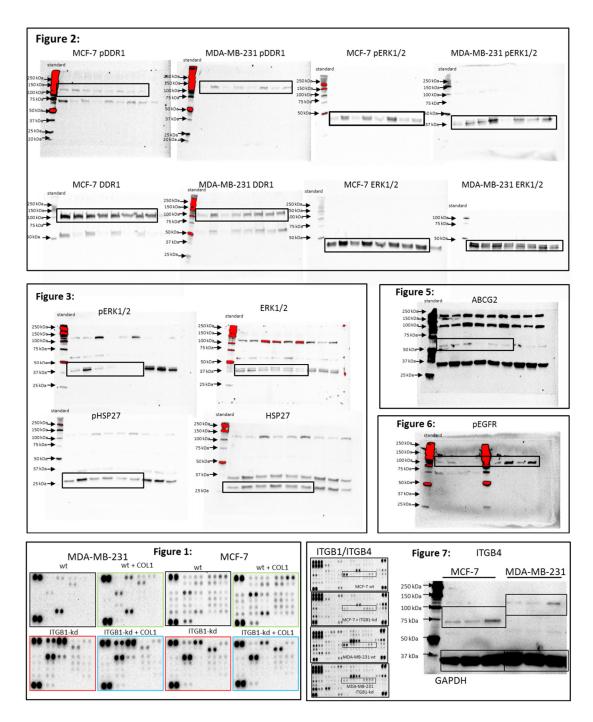


Figure S5: Representative uncut Western blots and proteome profiler arrays, which are partly shown in Figures 1–7.



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