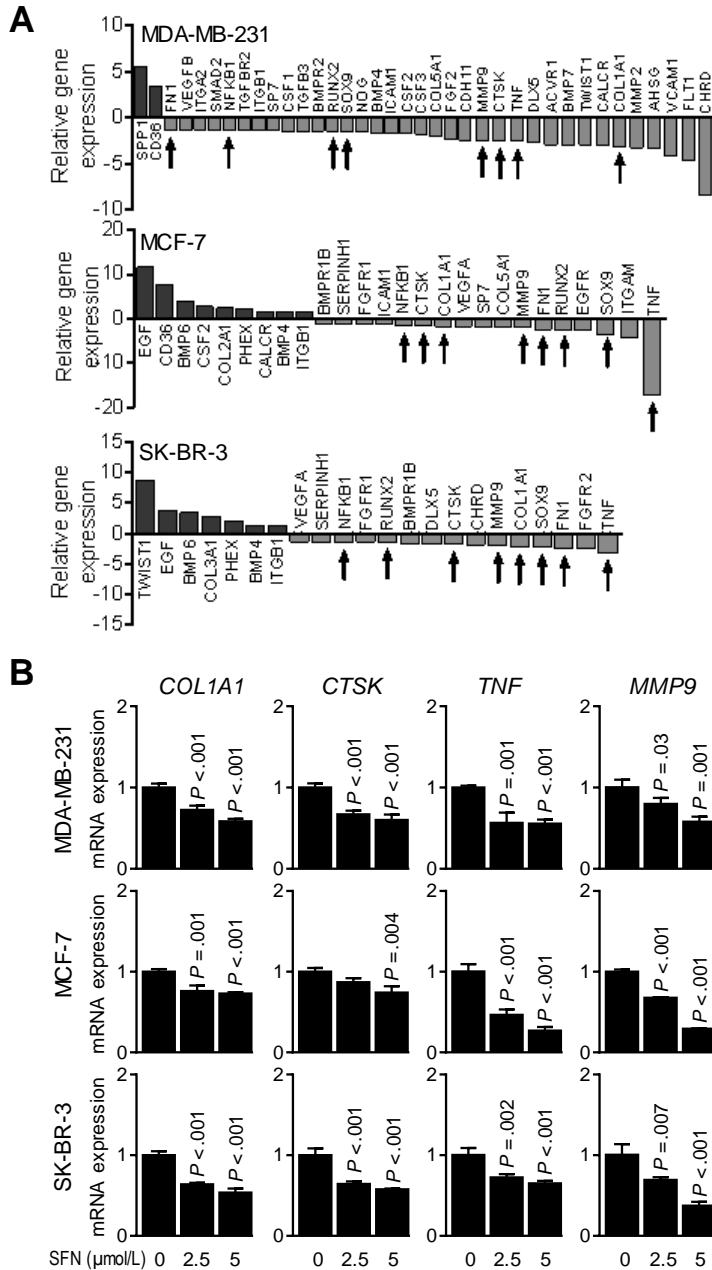
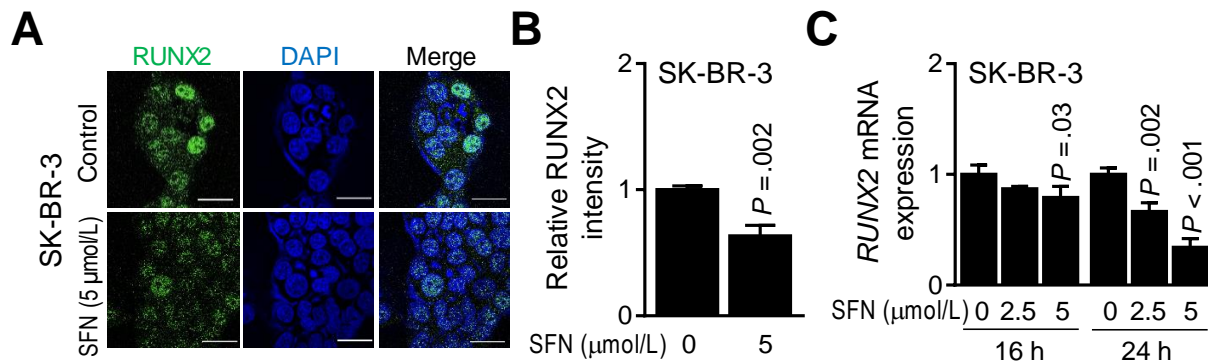


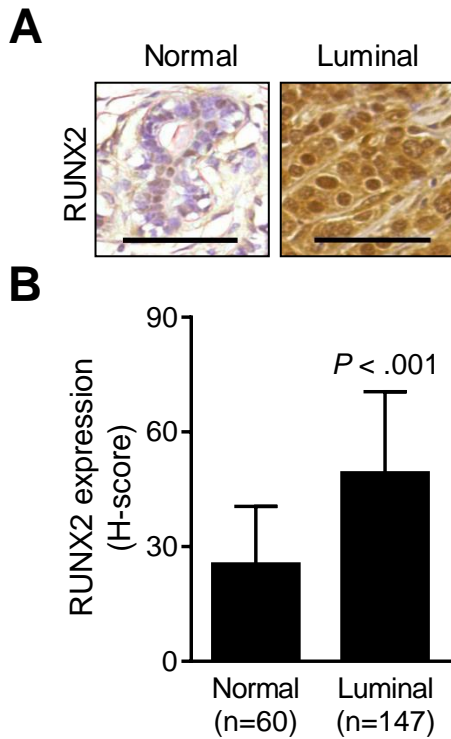
Supplementary Figure S1: Summary of gene expression changes following sulforaphane (SFN) treatment in breast cancer cells. **A**, Relative gene expression in SFN-treated cells in comparison with control. Arrows indicate common down-regulated genes in all three cell lines. **B**, Quantitative reverse transcription-polymerase chain reaction (qRT-PCR) for *collagen, type I, alpha 1 (COL1A1)*, *cathepsin K (CTSK)*, *tumor necrosis factor (TNF)*, and *matrix metalloproteinase (MMP9)* gene expression in control and SFN-treated cells (24-hour treatment). Results shown are mean \pm SD (n=3). Statistical significance of difference was analyzed by ANOVA followed by Dunnett's test. Consistent results were obtained from two independent experiments.



Supplementary Figure S2: Sulforaphane (SFN) treatment downregulated expression of Runt-related transcription factor 2 (RUNX2) in SK-BR-3 cell line. **A**, Representative confocal microscopic images ($\times 60$ objective magnification in oil) of RUNX2 protein (green) in SK-BR-3 cells after 24 hours of treatment with dimethyl sulfoxide (DMSO) or 5 $\mu\text{mol/L}$ SFN. Nuclei were stained with 4',6-diamidino-2-phenylindole (blue). **B**, Quantitation of RUNX2 protein level intensity in SK-BR-3 cells. Results shown are mean RUNX2 intensity \pm SD ($n=3$). Statistical significance of difference was analyzed by two-sided Student's t-test. **C**, Quantitative real-time PCR analysis of *RUNX2* mRNA expression in SK-BR-3 cells treated with DMSO or the indicated doses of SFN for specified time points. Results shown are mean \pm SD ($n=3$). Statistical significance of difference was analyzed by one-way analysis of variance (ANOVA) followed by Dunnett's test. Consistent results were obtained from repeated experiments.

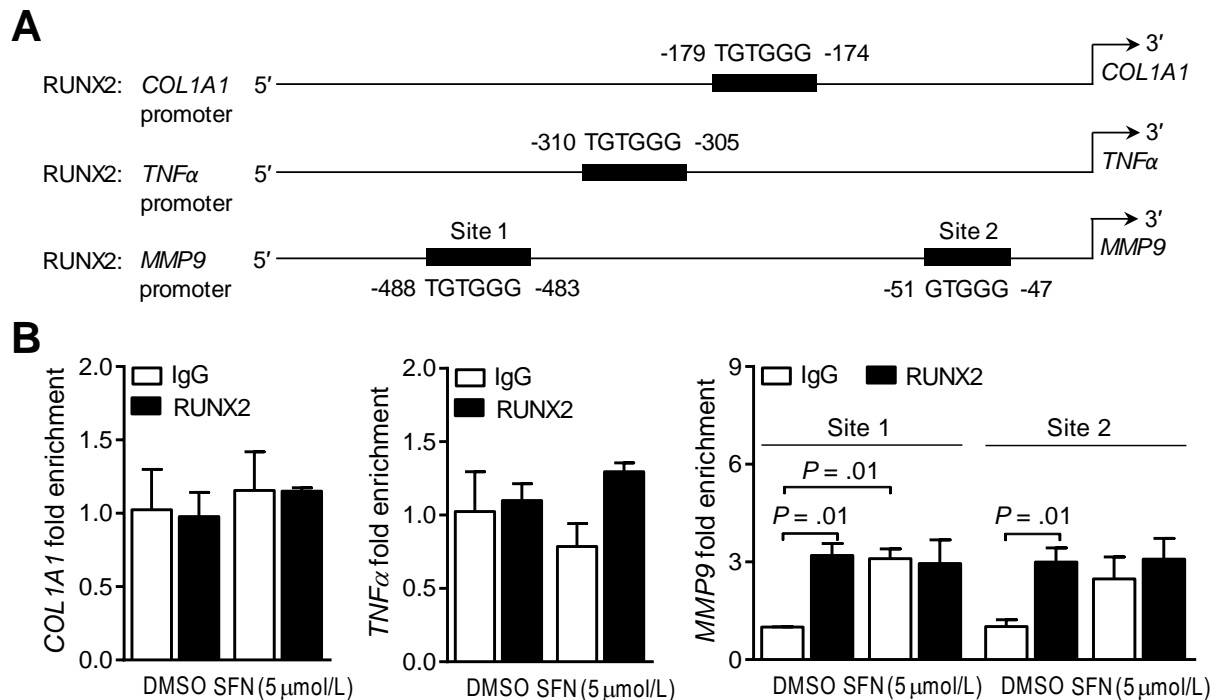


Supplementary Figure S3: Expression of runt-related transcription factor 2 (RUNX2) protein was higher in luminal-type breast cancers than in normal mammary tissues. **A**, Immunohistochemical images for RUNX2 protein (scale bar = 100 μm ; 20 \times objective magnification) in a representative normal mammary tissue and in a luminal-type breast cancer tissue. **B**, Quantitation (H-score) of RUNX2 protein level in normal mammary tissues (n = 60) and luminal-type breast cancer tissues (n = 147). Results shown are mean \pm SD. Statistical significance was determined by two-sided Student's t-test.



Supplementary Figure S4: Runt-related transcription factor 2 (RUNX2) was recruited at the promoter of *matrix metalloproteinase 9 (MMP9)* **A**, Putative RUNX2 binding sequence sites in the promoter region of *collagen, type I, alpha 1 (COL1A1)*, *tumor necrosis factor α (TNF α)*, and *matrix metalloproteinase (MMP9)*. The primers were as follows: *COL1A1*- Forward 5'-ATTGGAGGTCCCAGGAAGAG-3' Reverse 5'-TTGGGAGTTGGAATGGAGAG; *MMP9*- Primer for: -488 TGTGGG -483 Forward 5'-AGGCTGCTACTGTCCCCTTT-3' Reverse 5'-TCTGAAAGCCTCCAGTGGTC-3'; Primer for -51 GTGGG -47 Forward 5'-GAGTCAGCACTTGCCTGTCA-3' Reverse 5'-AAGAGCACAAGGGTGGACTG-3'; and *TNF α* - Forward 5'-GACAGATGTGGGGTGTGAGA-3' Reverse 5'-ACCTTCCAGGCATTCAACAG-3'. **B**, Chromatin immunoprecipitation assay using MDA-MB-231 cells showing the binding of RUNX2 transcription factor to the *MMP9*. Results shown are the mean \pm SD (n = 2-3). Statistical significance of difference was analyzed by Bonferroni's multiple comparisons test.

Supplementary Figure S4



Supplementary Table S1

Correlation between genes altered by SFN treatment (TCGA BrCa dataset, n = 1097).

The ns signifies not significant.

Gene	Gene	Pearson r	P
<i>RUNX2</i>	<i>NFKB1</i>	.22	<.001
<i>RUNX2</i>	<i>SOX9</i>	.059	.05
<i>RUNX2</i>	<i>CTSK</i>	.69	<.001
<i>RUNX2</i>	<i>COL1A1</i>	.68	<.001
<i>RUNX2</i>	<i>TNF</i>	.11	.004
<i>RUNX2</i>	<i>MMP9</i>	.24	<.001
<i>NFKB1</i>	<i>SOX9</i>	.08	.01
<i>NFKB1</i>	<i>CTSK</i>	.18	<.001
<i>NFKB1</i>	<i>COL1A1</i>	.11	<.001
<i>NFKB1</i>	<i>TNF</i>	.16	<.001
<i>NFKB1</i>	<i>MMP9</i>	.03	ns

Supplementary Table S2

Consensus binding sequences for transcription factors

Transcription factor	Target gene	No. of sites	Position
RUNX2	<i>NFKB1</i>	3	-1161 <i>TGTGGG</i> -1152; -901 <i>TGTGGG</i> -896; -793 <i>GTGGG</i> -789
RUNX2	<i>RELA</i>	4	-988 <i>AGTGGG</i> -983; -712 <i>TGTGGG</i> -707; -640 <i>GTGGG</i> -635; -322 <i>GTGGG</i> -318
RUNX2	<i>COL1A1</i>	1	-179 <i>TGTGGG</i> -174
RUNX2	<i>MMP9</i>	2	-488 <i>TGTGGG</i> -483; -51 <i>GTGGG</i> -47
RUNX2	<i>TNF</i>	1	-310 <i>TGTGGG</i> -305
RUNX2	<i>SOX9, CTSK</i>	0	No match
NFKB1	<i>RELA</i>	5	-1054 <i>TCCCC</i> -1050; -798 <i>GGGGAT</i> -793; -780 <i>GGGGGTxxxxxxxCCCC</i> -764; -729 <i>GGGGGA</i> -724; -350 <i>ATTCCCC</i> -344
NFKB1	<i>COL1A1</i>	5	-902 <i>GGGGGAT</i> -896; -886 <i>GGGGAT</i> -881; -848 <i>GAATCCCC</i> -841; -588 <i>GTTCCCC</i> -582; -416 <i>GGAGGACCCCC</i> -406
NFKB1	<i>MMP9</i>	2	-619 <i>GGGGGTTGCCCC</i> -608; -332 <i>GGGGGATCCC</i> -323
NFKB1	<i>RUNX2, SOX9, CTSK, TNF</i>	0	No match

Consensus binding sequences according to JASPAR (<http://jaspar.genereg.net/>) browser:

RUNX2: **TGTGGTTT**; NF- κ B1: **GGGGATTCCCC**