

# PNAS

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Supplementary Information for

*Loss of BOP1 confers resistance to BRAF kinase inhibitors in melanoma by activating MAP kinase pathway*

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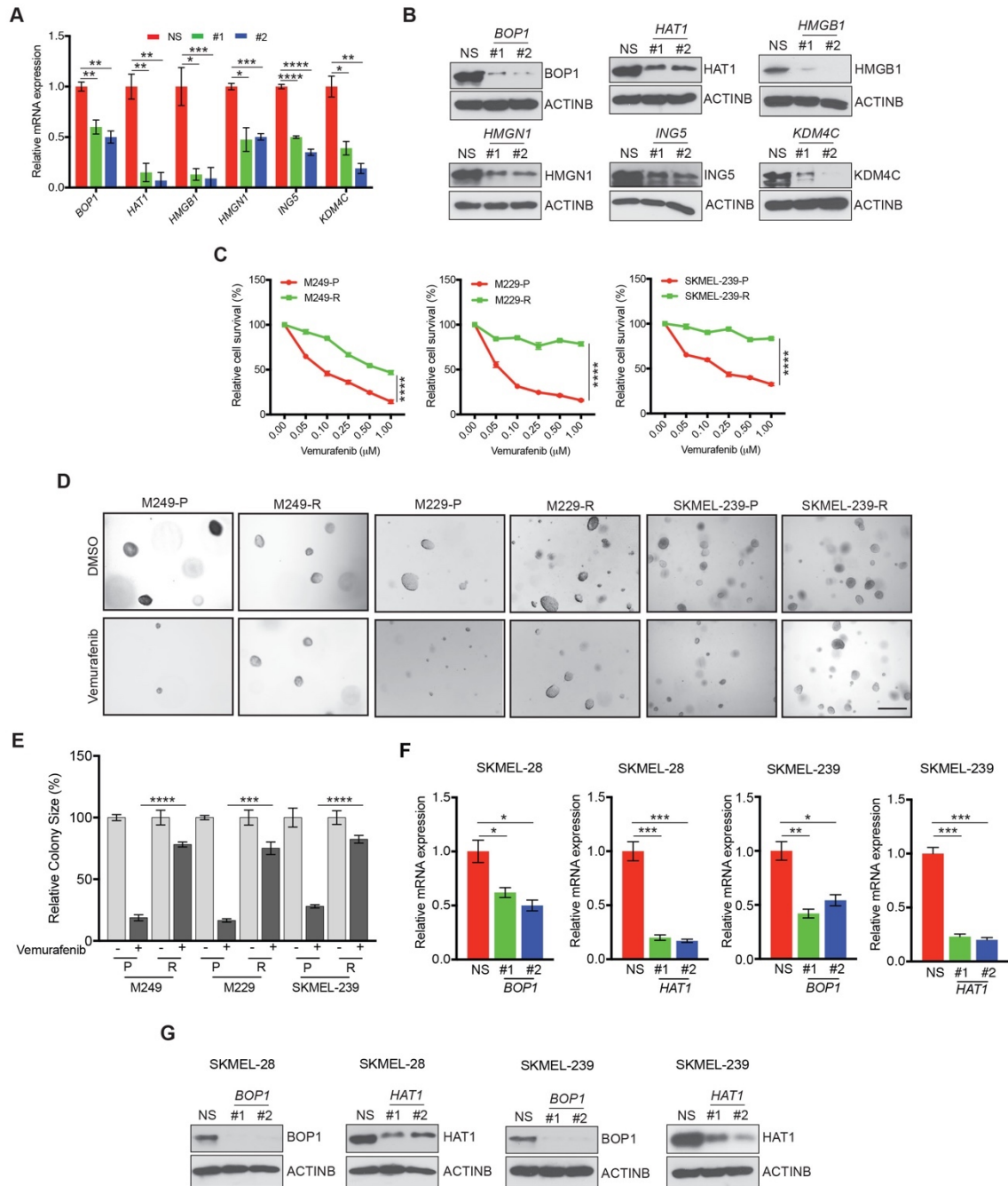
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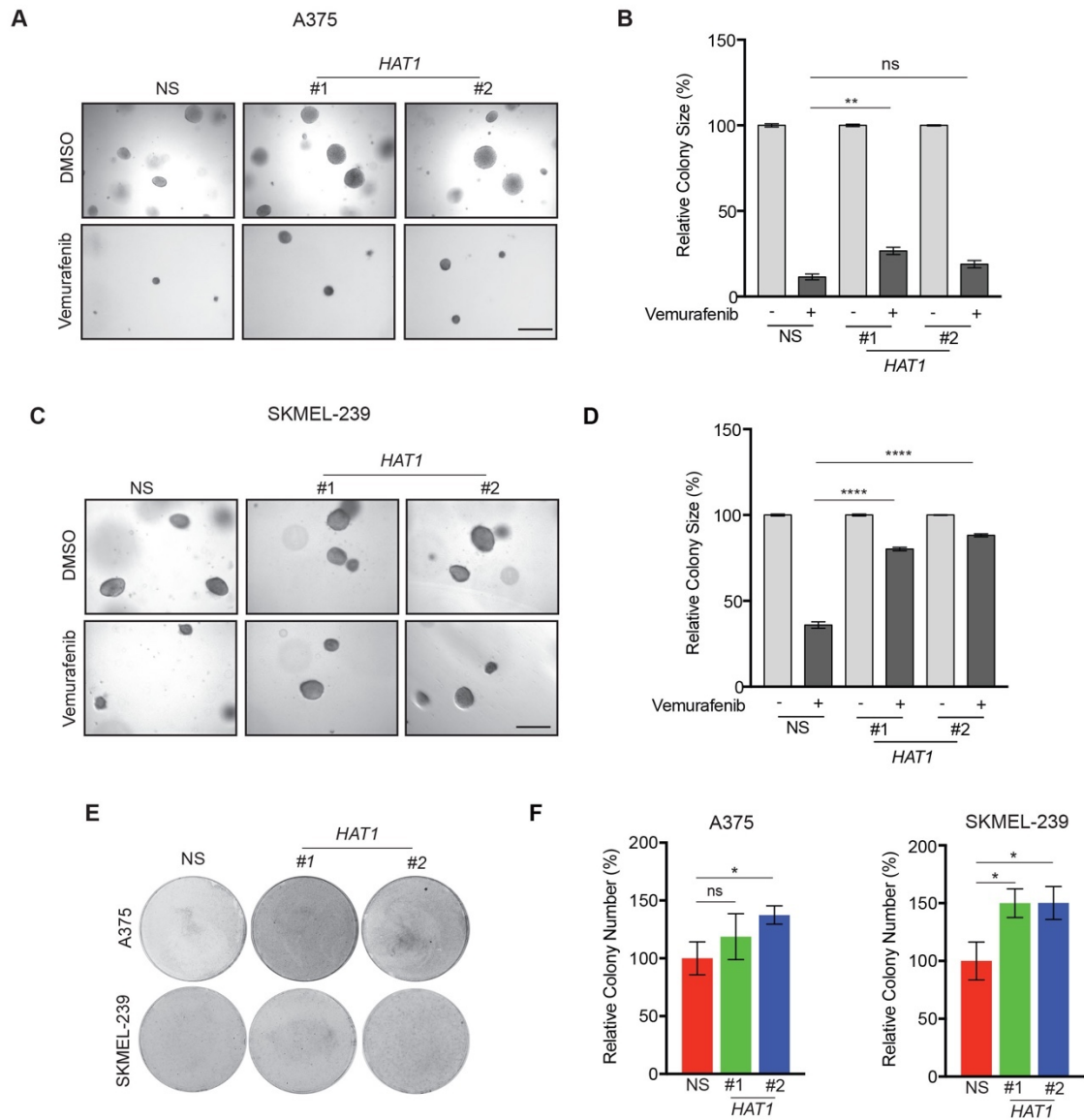
**Fig. S1.**



**Fig. S1. Short hairpin RNA (shRNA) targeting efficiencies and vemurafenib sensitivity of parental and BRAF kinase inhibitor-resistant melanoma cells.** (A) Expression of the indicated genes measured by RT-qPCR in A375 cells expressing non-specific (NS) or indicated shRNAs; mRNA expression in shRNA-targeted cells relative to NS shRNA is shown. (B) Levels of indicated proteins measured by immunoblotting in A375 cells expressing NS or shRNAs. (C) Relative cell survival (%) measured by MTT assay of indicated parental (P) or BRAF kinase inhibitor (BRAFi)-resistant (R) melanoma cells treated for 3 days with the indicated concentrations of vemurafenib or

DMSO control. (D) Indicated parental (P) or BRAFi-resistant (R) melanoma cells were treated with vemurafenib (1  $\mu$ M) or DMSO control, and anchorage-independent growth was assessed by soft-agar assay. Representative images of soft-agar colonies under the indicated conditions are shown. Scale bar, 500  $\mu$ M. (E) Relative colony size (%) for the soft-agar data in panel D. (F) Expression of the indicated genes measured by RT-qPCR in SKMEL-28 or SKMEL-239 cells expressing a NS or shRNAs targeting the indicated genes; mRNA expression in shRNA-targeted cells relative to NS shRNA is shown. (G) Levels of indicated proteins measured by immunoblotting in SKMEL-28 or SKMEL-239 cells expressing NS or shRNAs against indicated genes. ACTINB was used as a loading control. Data are presented as the mean  $\pm$  standard error of the mean (SEM). \* $P$ <0.05, \*\* $P$ <0.01, \*\*\* $P$ <0.001, and \*\*\*\* $P$ <0.0001.

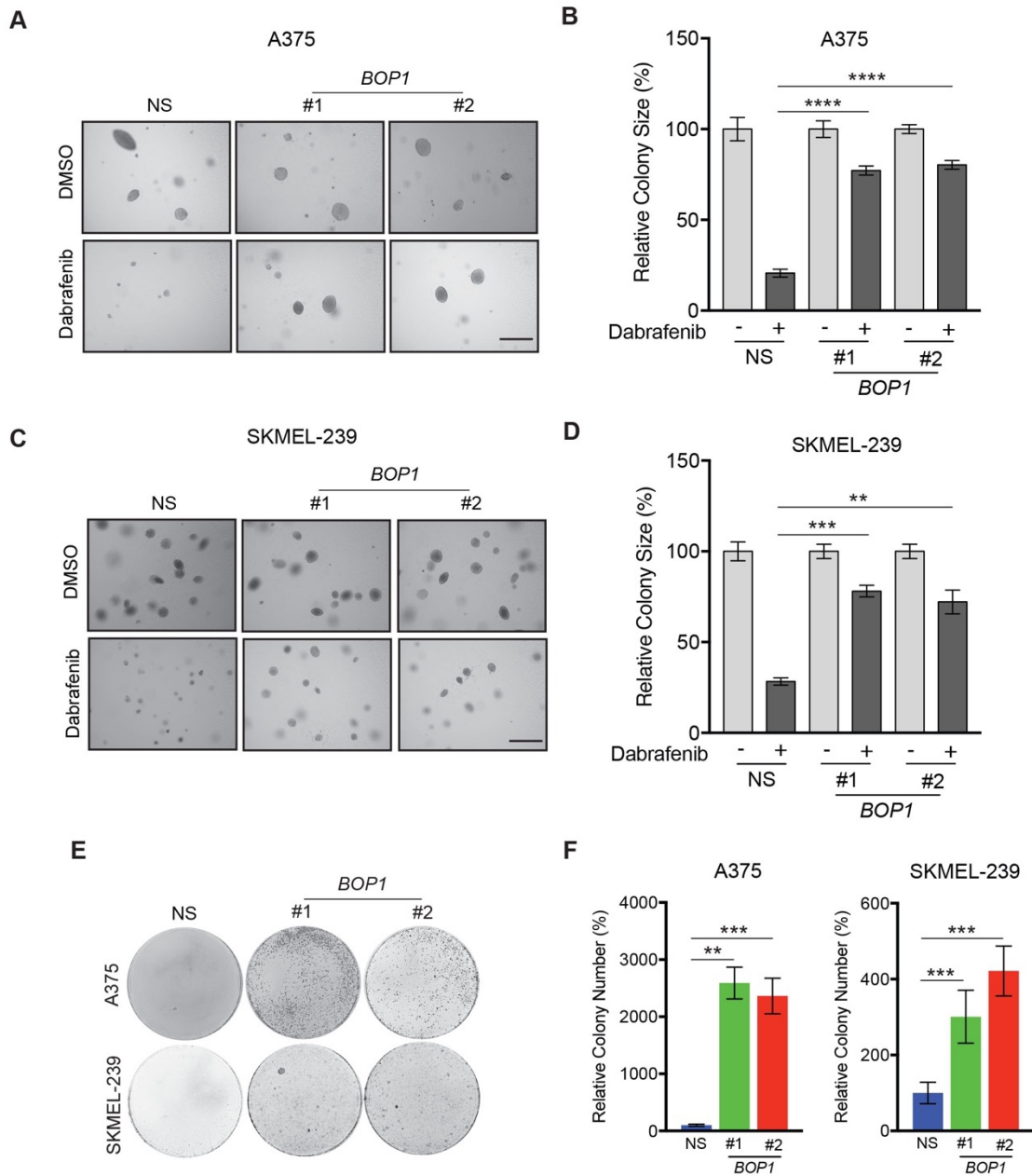
**Fig. S2.**



**Fig. S2. *HAT1* knockdown confers vemurafenib resistance in BRAF-mutant melanoma cells.** (A) A375 cells expressing non-specific (NS) or *HAT1* shRNAs were treated with vemurafenib (1  $\mu$ M) or DMSO control, and anchorage-independent growth was measured by soft-agar assay. Representative images of soft-agar colonies under the indicated conditions are shown. Scale bar, 500  $\mu$ M. (B) Relative colony size (%) for the data in panel A. (C) SKMEL-239 cells expressing NS or *HAT1* shRNAs were treated with vemurafenib (1  $\mu$ M) or DMSO control, and anchorage-independent growth was measured by soft-agar assay. Representative images of soft-agar colonies under the indicated conditions are shown. Scale bar, 500  $\mu$ M. (D) Relative colony size (%) for the data in panel C. (E) A375 and SKMEL-239 cells expressing NS or *HAT1* shRNAs were treated with vemurafenib (2  $\mu$ M) and analysed by clonogenic assay. Representative clonogenic assay plates are shown. (F) Relative colony number (%) for the experiments

in panel E. Data are presented as the mean  $\pm$  SEM. ns = not significant.  
\* $P < 0.05$ , \*\* $P < 0.001$ , and \*\*\*\* $P < 0.0001$ .

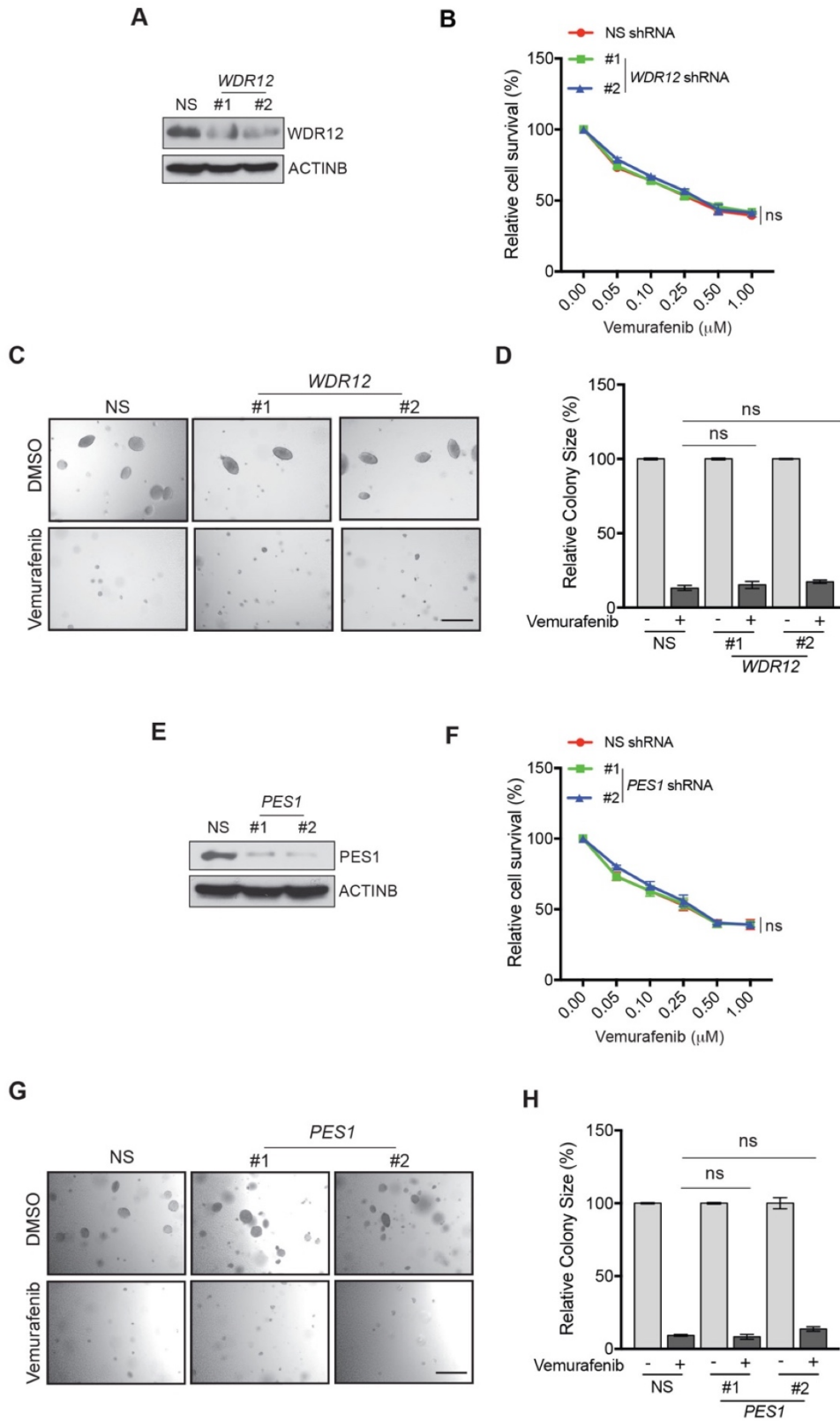
**Fig. S3.**



**Fig. S3. Knockdown of *BOP1* confers resistance to the BRAFi, dabrafenib.** (A) A375 cells expressing non-specific (NS) or *BOP1* shRNAs were treated with dabrafenib (50 nM) or DMSO control, and anchorage-independent growth was measured by soft-agar assay. Representative images of soft-agar colonies under the indicated conditions are shown. Scale bar, 500  $\mu$ M. (B) Relative colony size (%) for the data in panel A. (C) SKMEL-239 cells expressing NS or *BOP1* shRNAs were treated with dabrafenib (50 nM) or DMSO control, and anchorage-independent growth was measured by soft-agar assay. Representative images of soft-agar colonies under the indicated conditions are shown. Scale bar, 500  $\mu$ M. (D) Relative colony size (%) for the data in panel C. (E) A375 and SKMEL-239 cells expressing NS or *BOP1* shRNAs were treated with dabrafenib (100 nM) and analysed by clonogenic assay. Representative clonogenic assay plates are

shown. (F) Relative colony number (%) for the data in panel E. Data are presented as the mean  $\pm$  SEM. \*\* $P < 0.01$ , \*\*\* $P < 0.001$ , and \*\*\*\* $P < 0.0001$ .

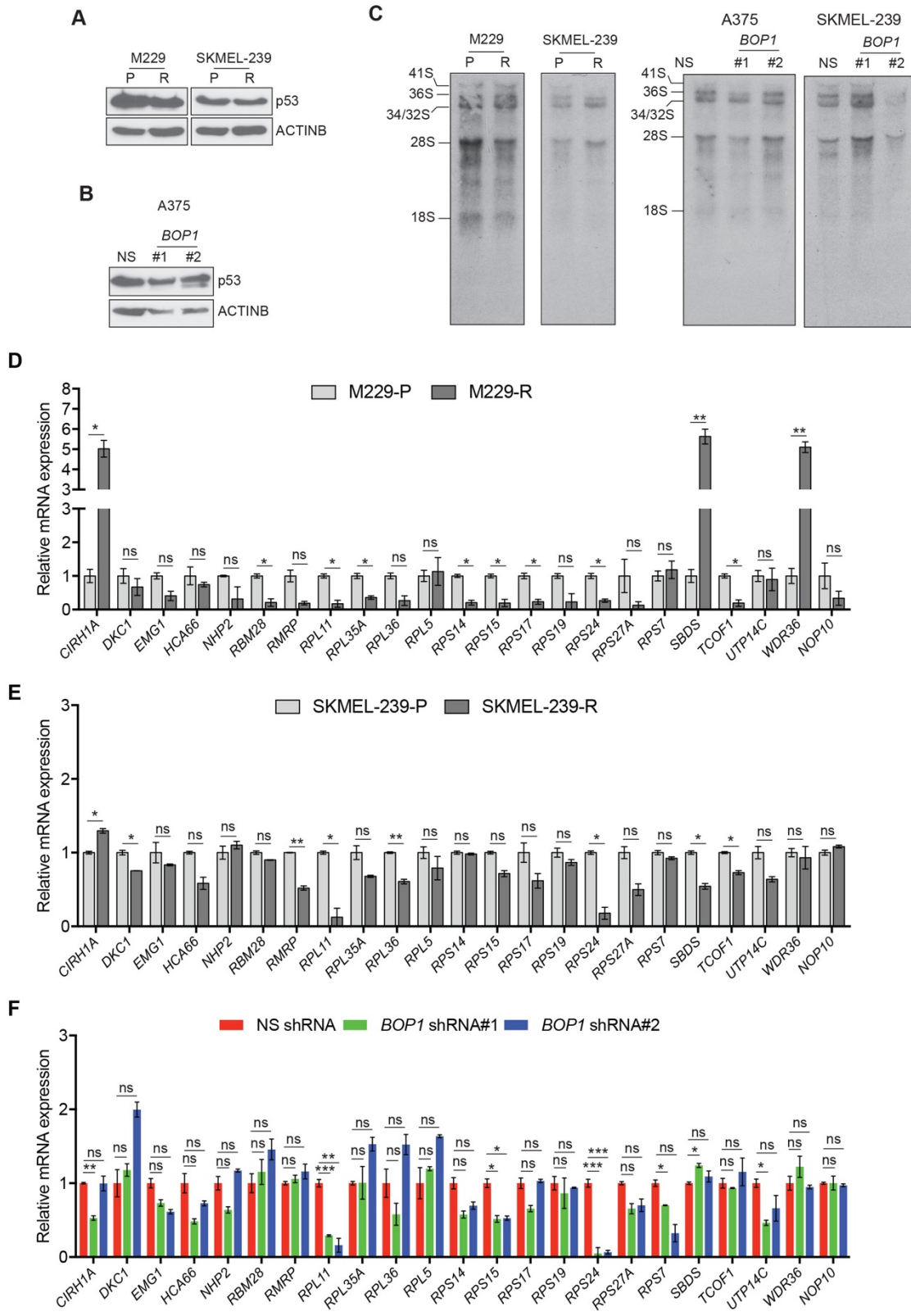
**Fig. S4.**





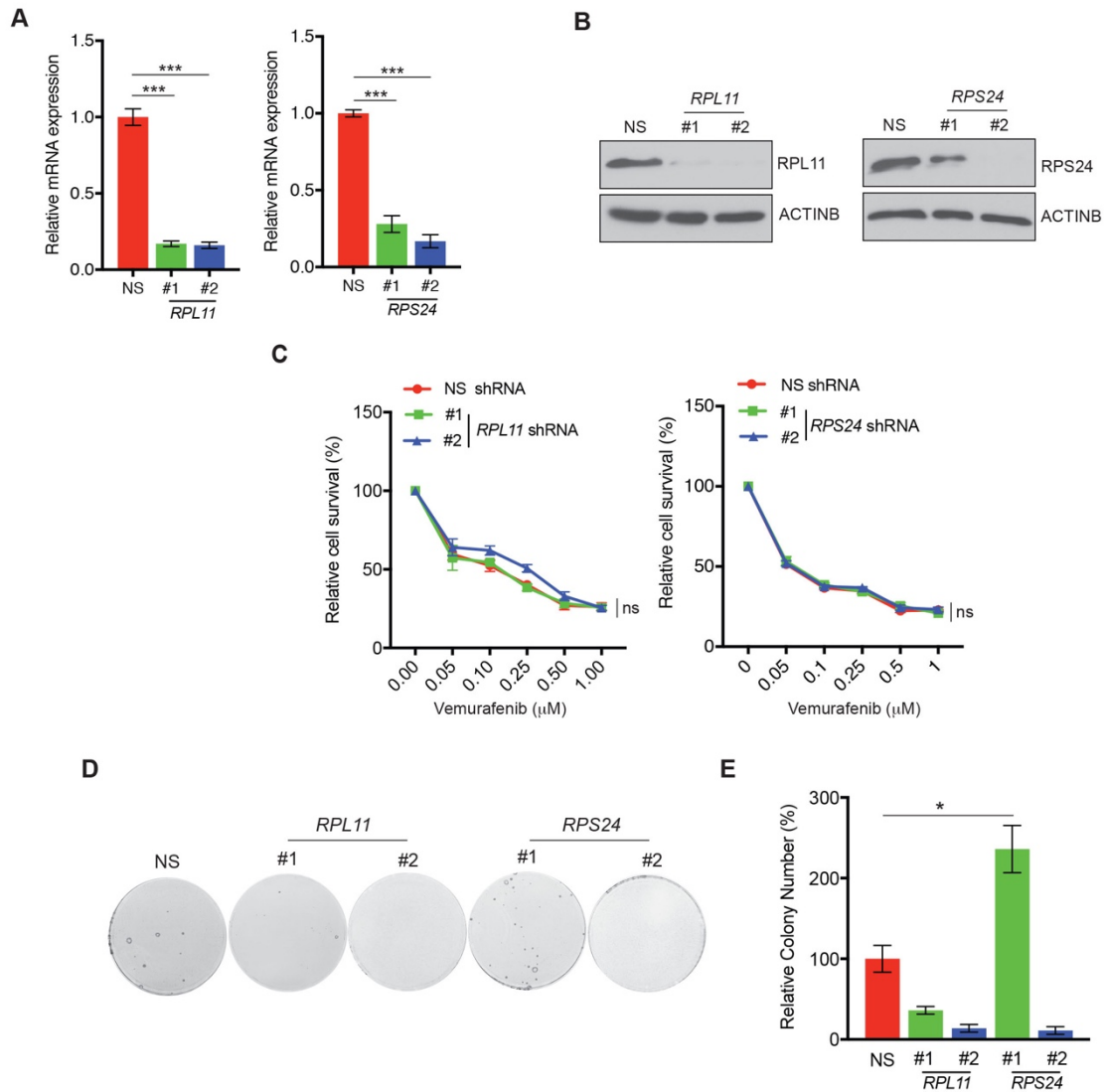
**Fig. S4. Knockdown of the PES1-BOP1-WDR12 complex proteins, WDR12 and PES1, does not confer resistance to BRAF inhibitors.** (A) Levels of indicated proteins measured by immunoblotting in A375 cells expressing non-specific (NS) or *WDR12* shRNAs. (B) Relative cell survival (%) measured by MTT assay in A375 cells expressing NS or *WDR12* shRNAs and treated with indicated concentrations of vemurafenib or DMSO control for 3 days. (C) A375 cells expressing NS or *WDR12* shRNAs were treated with vemurafenib (1  $\mu$ M) or DMSO control, and anchorage-independent growth was measured by soft-agar assay. Representative images of soft-agar colonies under the indicated conditions are shown. Scale bar, 500  $\mu$ M. (D) Relative colony size (%) for the data in panel C. (E) Levels of indicated proteins measured by immunoblotting in A375 cells expressing NS or *PES1* shRNAs. (F) Relative cell survival (%) measured by MTT assay in A375 cells expressing NS or *PES1* shRNAs and treated with indicated concentrations of vemurafenib or DMSO control for 3 days. (G) A375 cells expressing NS or *PES1* shRNAs were treated with vemurafenib (1  $\mu$ M) or DMSO control, and anchorage-independent growth was measured by soft-agar assay. Representative images of soft-agar colonies under the indicated conditions are shown. Scale bar, 500  $\mu$ M. (H) Relative colony size (%) for the data in panel G. Data are presented as the mean  $\pm$  SEM. ns = not significant.

**Fig. S5.** Type or paste caption here. Paste figure above the caption.



**Fig. S5. Ribosomopathy does not drive BRAFi resistance.** (A) Protein levels of p53 and ACTINB were measured by immunoblotting in the indicated BRAFi-sensitive parental (P) and BRAFi-resistant (R) cell lines. (B) Protein levels of p53 and ACTINB were measured by immunoblotting in A375 cells expressing non-specific (NS) or *BOPI* shRNAs. (C) Ribosomal RNA (rRNA) biogenesis in (Left) indicated BRAFi-sensitive parental (P) and resistant (R) cells and (Right) A375 cells expressing NS or *BOPI* shRNAs, as measured by tritiated uridine ( $H^3$ -U)-based metabolic labelling. (D) Expression of the indicated genes was measured in BRAFi-sensitive parental (P) and BRAFi-resistant (R) M229 cells by RT-qPCR; mRNA expression in R cells relative to P cells is shown. (E) Expression of the indicated genes was measured in BRAFi-sensitive parental (P) and BRAFi-resistant (R) SKMEL-239 cells by RT-qPCR; mRNA expression in R cells relative to P cells is shown. (F) Expression of the indicated genes was measured in A375 cells expressing NS or *BOPI* shRNAs by RT-qPCR; mRNA expression in shRNA-targeted cells relative to NS shRNA is shown. Data are presented as the mean  $\pm$  SEM. ns = not significant, \* $P$ <0.05, \*\* $P$ <0.01 and \*\*\* $P$ <0.001.

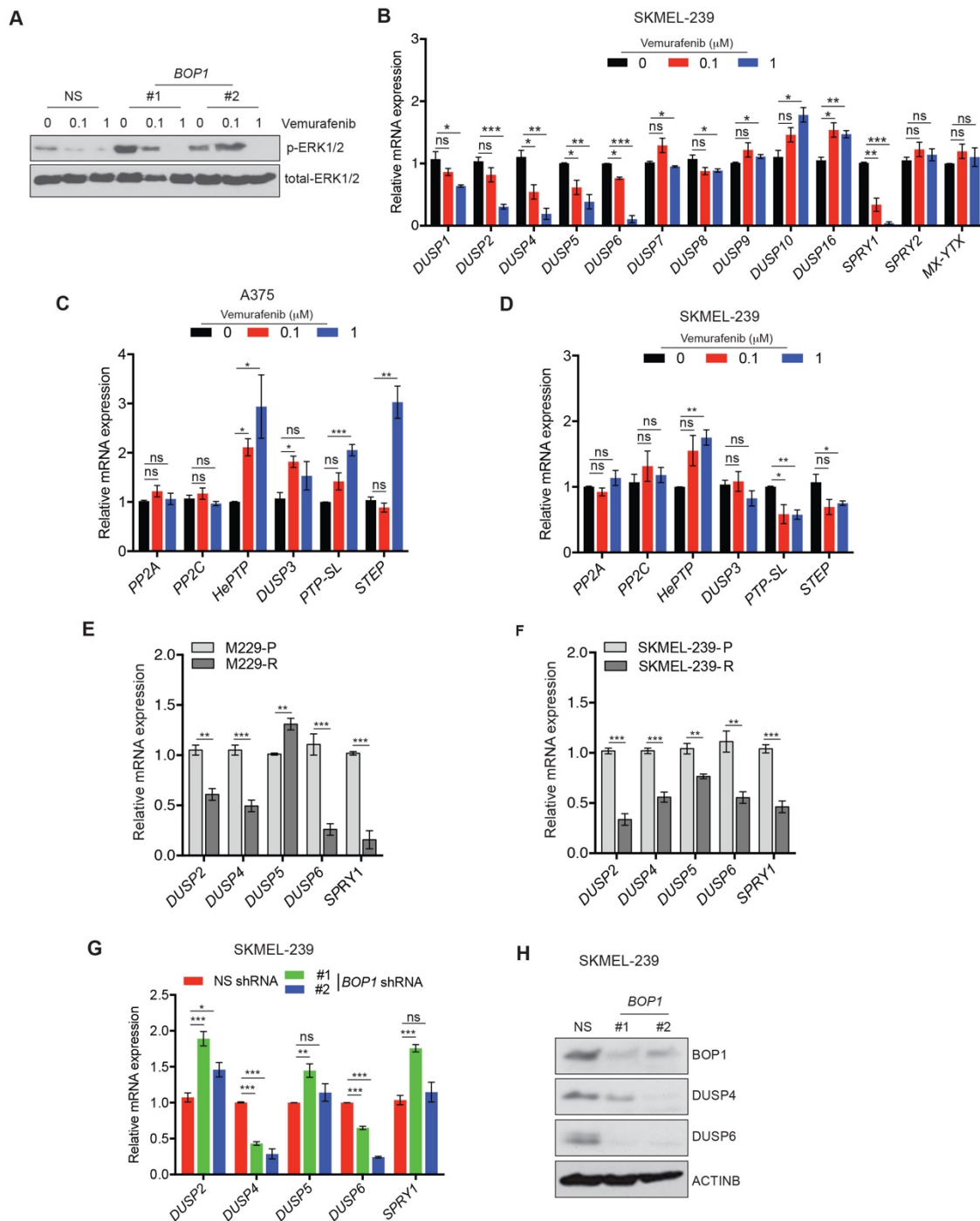
**Fig. S6.** Type or paste caption here. Paste figure above the caption.



**Fig. S6. Loss of ribosomopathy genes does not confer resistance to BRAF inhibitor.**

(A) Expression of *RPL11* and *RPS24* was measured by RT-qPCR in A375 cells containing non-specific (NS), *RPL11*, or *RPS24* shRNAs, as indicated; mRNA expression in shRNA-targeted cells relative to NS shRNA is shown. (B) Protein levels of RPL11 or RPS24 were measured by immunoblotting in A375 cells containing NS, *RPL11*, or *RPS24* shRNAs, as indicated. ACTINB was used as a loading control. (C) Relative cell survival (%) measured by MTT assay in A375 cells expressing NS, *RPL11*, or *RPS24* shRNAs that were treated with indicated concentrations of vemurafenib for 3 days. (D) A375 cells expressing NS, *RPL11*, or *RPS24* shRNAs were treated with vemurafenib (2  $\mu$ M) or DMSO control and analysed by clonogenic assay. Representative clonogenic assay plates are shown. (E) Relative colony numbers (%) for the experiments in panel D. Data are presented as the mean  $\pm$  SEM. ns = not significant. \* $P$ <0.05 and \*\*\* $P$ <0.001.

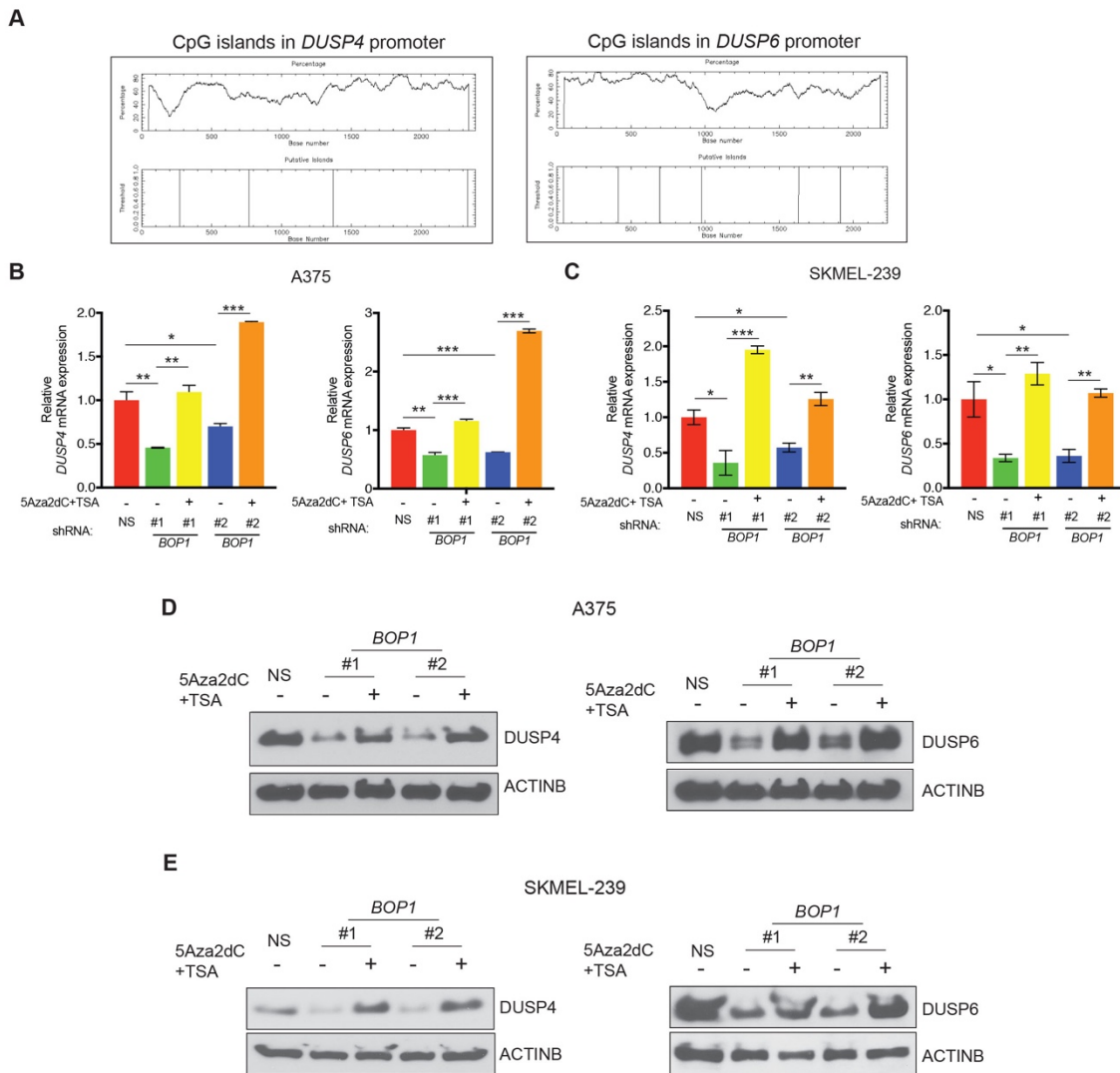
**Fig. S7.**



**Fig. S7. Loss of BOP1 enhances the MAP kinase pathway by regulating *DUSP4* and *DUSP6*.** (A) Levels of indicated proteins were measured by immunoblotting in SKMEL-239 cells expressing non-specific (NS) or *BOP1* shRNAs and treated with indicated concentrations of vemurafenib (0.1 or 1.0 μM) or DMSO for 3 h. (B) Levels of indicated phosphatases were measured by RT-qPCR in SKMEL-239 cells treated with vemurafenib (0.1 or 1 μM) or DMSO control for 24 h. Relative mRNA expression of indicated phosphatases in vemurafenib-treated relative to DMSO-treated cells is shown. (C) Levels

of indicated phosphatases were measured by RT-qPCR in A375 cells treated with vemurafenib (0.1 or 1  $\mu$ M) or DMSO control for 24 h. Relative mRNA expression of indicated phosphatases in vemurafenib-treated relative to DMSO-treated cells is shown. (D) Levels of indicated phosphatases were measured by RT-qPCR in SKMEL-239 cells treated with vemurafenib (0.1 or 1  $\mu$ M) or DMSO control for 24 h. Relative mRNA expression of indicated phosphatases in vemurafenib-treated relative to DMSO-treated cells is shown. (E) Levels of indicated phosphatases were measured by RT-qPCR in M229 parent (P) or BRAFi-resistant M229 (M229-R) cells. Relative mRNA expression of indicated phosphatases in M229-R relative to P cells is shown. (F) Levels of indicated phosphatases were measured by RT-qPCR in SKMEL-239 parent (P) or BRAFi-resistant SKMEL-239 (SKMEL-239-R) cells. Relative mRNA expression of indicated phosphatases in SKMEL-239-R relative to P cells is shown. (G) Levels of indicated phosphatases were measured by RT-qPCR in SKMEL-239 cells expressing NS or *BOP1* shRNAs. Relative mRNA expression in shRNA-targeted cells relative to NS shRNA is shown. (H) Levels of indicated proteins were measured by immunoblotting in SKMEL-239 cells expressing NS or *BOP1* shRNAs. Data are presented as the mean  $\pm$  SEM. ns = not significant. \* $P$ <0.05, \*\* $P$ <0.01, and \*\*\* $P$ <0.001.

**Fig. S8.**



**Fig. S8. BOP1 loss results in transcriptional silencing of *DUSP4* and *DUSP6*.** (A) The European Bioinformatic Institute (EBI) CpG plot analysis of the promoter sequences for *DUSP4* and *DUSP6* reveal the presence of CpG islands. (B) Expression of *DUSP4* and *DUSP6* was measured by RT-qPCR in A375 cells containing non-specific (NS) or *BOP1* shRNAs, treated with 5Aza2dC (5  $\mu$ M; 72 h) and TSA (1  $\mu$ M; 12 h). Relative mRNA expression in shRNA-targeted cells relative to NS shRNA is shown. (C) Expression of *DUSP4* and *DUSP6* was measured by RT-qPCR in SKMEL-239 cells expressing NS or *BOP1* shRNAs, treated with 5Aza2dC (5  $\mu$ M; 72 h) and TSA (1  $\mu$ M; 12 h). Relative mRNA expression in shRNA-targeted cells relative to NS shRNA is shown. (D) Levels of DUSP4 and DUSP6 proteins were measured by immunoblotting in A375 cells expressing NS or *BOP1* shRNAs, treated with 5Aza2dC (5  $\mu$ M, 72 h) and TSA (1  $\mu$ M, 12 h). ACTINB was used as a loading control. (E) Levels of DUSP4 and DUSP6 proteins were measured by immunoblotting in SKMEL-239 cells expressing NS or *BOP1* shRNAs, treated with 5Aza2dC (5  $\mu$ M, 72 h) and TSA (1  $\mu$ M, 12 h). ACTINB was used

as a loading control. Data are presented as the mean  $\pm$  SEM. \* $P$ <0.05, \*\* $P$ <0.01, and \*\*\* $P$ <0.001.



**Table S1.** List of genes targeted by the human epigenome-wide shRNA library and shRNA IDs.

S.No.	Gene Name	shRNA IDs
1	2410016006Rik	TRCN0000180487
2	ACTL6A	TRCN0000072273 TRCN0000072274 TRCN0000072275 TRCN0000072277 TRCN0000072276
3	ACTL6B	TRCN0000116767 TRCN0000116768 TRCN0000116769 TRCN0000116770 TRCN0000116771
4	AIRE	TRCN0000018890 TRCN0000018891 TRCN0000018892 TRCN0000018893 TRCN0000018894
5	ALKBH2	TRCN0000064678 TRCN0000064679 TRCN0000064680 TRCN0000064681 TRCN0000064682
6	AOF1/KDM1B	TRCN0000046073 TRCN0000046074 TRCN0000046076 TRCN0000046075 TRCN0000046077
7	AOF2	TRCN0000046068 TRCN0000046069 TRCN0000046070 TRCN0000046071 TRCN0000046072
8	APITD1	TRCN0000038878 TRCN0000038874 TRCN0000191787

9	ARID1A	TRCN0000059089 TRCN0000059090 TRCN0000059092 TRCN0000059088 TRCN0000059091 TRCN0000071394 TRCN0000071397
10	ARID3C	TRCN0000201971
11	ARID4A	TRCN0000014595 TRCN0000014593 TRCN0000014594 TRCN0000014596 TRCN0000014597
12	ASF1A	TRCN0000074268 TRCN0000074269 TRCN0000074271 TRCN0000074272 TRCN0000074270
13	ASF1B	TRCN0000074225 TRCN0000074226 TRCN0000074227 TRCN0000108962
14	ASH1L	TRCN0000016168 TRCN0000016169 TRCN0000016170 TRCN0000016171 TRCN0000016172
15	ASH2L	TRCN0000019274 TRCN0000019275 TRCN0000019276 TRCN0000019277 TRCN0000019278
16	ASXL1	TRCN0000135296 TRCN0000138572 TRCN0000133699 TRCN0000134690 TRCN0000135579
17	ATM	TRCN0000010299 TRCN0000038654 TRCN0000039948

18	ATR	TRCN0000039614 TRCN0000007749 TRCN0000023912 TRCN0000023913
19	ATRX	TRCN0000013588 TRCN0000013589 TRCN0000013590 TRCN0000013591 TRCN0000013592
20	AURKB	TRCN0000000776 TRCN0000010547 TRCN0000000777 TRCN0000000778 TRCN0000000779
21	BAZ1A	TRCN0000034279 TRCN0000034280 TRCN0000034281 TRCN0000034282 TRCN0000034283
22	BAZ1B	TRCN0000013339 TRCN0000013342 TRCN0000013340 TRCN0000013341 TRCN0000013338 TRCN0000108939
23	BAZ2A	TRCN0000015568 TRCN0000015569 TRCN0000015571 TRCN0000015572 TRCN0000015570
24	BAZ2B	TRCN0000019928 TRCN0000019924 TRCN0000019926 TRCN0000019925 TRCN0000019927 TRCN0000116909 TRCN0000116910 TRCN0000116908 TRCN0000200866 TRCN0000191572

		TRCN0000201016
25	BCOR	TRCN0000033459 TRCN0000033460 TRCN0000033461 TRCN0000033462 TRCN0000033463
26	BMI1	TRCN0000020154 TRCN0000020155 TRCN0000020156 TRCN0000020157 TRCN0000020158
27	BOP1	TRCN0000078068 TRCN0000078069 TRCN0000078070 TRCN0000078071 TRCN0000078072
28	BPTF	TRCN0000016818 TRCN0000016819 TRCN0000016820 TRCN0000016821 TRCN0000016822
29	BRAF	TRCN0000006289 TRCN0000006290 TRCN0000006291 TRCN0000006292 TRCN0000006293
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		TRCN0000021378
33	BRD4	TRCN0000021424 TRCN0000021425 TRCN0000021426 TRCN0000021427 TRCN0000021428
34	BRD7	TRCN0000151378 TRCN0000151543 TRCN0000154102
35	BRD8	TRCN0000019379 TRCN0000019380 TRCN0000019381 TRCN0000019382 TRCN0000019383 TRCN0000096646
36	BRD9	TRCN0000127634 TRCN0000127780 TRCN0000128333 TRCN0000128610 TRCN0000131081
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		TRCN0000062667 TRCN0000062664 TRCN0000062665 TRCN0000062666 TRCN0000084534
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44	C20orf20/MRGBP	TRCN0000158734 TRCN0000159338 TRCN0000159786 TRCN0000159916 TRCN0000159917
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53	CBX8	TRCN0000021894 TRCN0000021895 TRCN0000021896 TRCN0000021897 TRCN0000021898
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70	CLOCK	TRCN0000018974

		TRCN0000018975 TRCN0000018976 TRCN0000018977 TRCN0000018978
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88	EHMT1	TRCN0000036054 TRCN0000036055 TRCN0000036056 TRCN0000036057 TRCN0000036058
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116	HDAC10	TRCN0000004859 TRCN0000004860 TRCN0000004861 TRCN0000004862 TRCN0000004863
117	HDAC11	TRCN0000017753 TRCN0000017754 TRCN0000017755 TRCN0000017756 TRCN0000017757
118	HDAC2	TRCN0000004819 TRCN0000004820 TRCN0000004821 TRCN0000004822 TRCN0000004823
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121	HDAC5	TRCN0000004834 TRCN0000004835 TRCN0000004836 TRCN0000004837 TRCN0000004838
122	HDAC6	TRCN0000004839 TRCN0000004840 TRCN0000004841 TRCN0000004842 TRCN0000004843
123	HDAC7	TRCN0000004844



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124	HDAC8	TRCN0000004849 TRCN0000004850 TRCN0000004851 TRCN0000004852 TRCN0000004853
125	HDAC9	TRCN0000004854 TRCN0000004855 TRCN0000004856 TRCN0000004857 TRCN0000004858
126	HELLS	TRCN0000000306 TRCN0000000307 TRCN0000000308 TRCN0000000309 TRCN0000000310
127	HILS1	TRCN0000106970 TRCN0000106971 TRCN0000106972 TRCN0000106973 TRCN0000106974
128	HIRA	TRCN0000020516 TRCN0000020515 TRCN0000020514 TRCN0000020515 TRCN0000020516 TRCN0000020517 TRCN0000020518
129	HLTF	TRCN0000003971 TRCN0000003972 TRCN0000003973 TRCN0000010836 TRCN0000003974
130	HMG20B	TRCN0000015500 TRCN0000015499 TRCN0000015498 TRCN0000015501

131	HMGA1	TRCN000004909 TRCN0000018953 TRCN0000018950 TRCN0000018951 TRCN0000018949 TRCN0000018952
132	HMGA2	TRCN0000021964 TRCN0000021967 TRCN0000021968 TRCN0000021965 TRCN00000126047 TRCN0000021966
133	HMGB1	TRCN0000018930 TRCN0000018932 TRCN0000018934 TRCN0000092693 TRCN0000018933 TRCN0000063714 TRCN0000018931
134	HMGB2	TRCN0000019009 TRCN0000019010 TRCN0000019013 TRCN0000019012 TRCN0000019011 TRCN0000092929 TRCN0000092932 TRCN0000092942
135	HMGB3	TRCN0000018518 TRCN0000018519 TRCN0000018520 TRCN0000071432 TRCN0000016705
136	HMGN1	TRCN0000019059 TRCN0000019060 TRCN0000019061 TRCN0000019062 TRCN0000019063
137	HMGN2	TRCN0000018551 TRCN0000018549 TRCN00000127823

		TRCN0000018552 TRCN0000108988 TRCN0000130247 TRCN0000018548 TRCN0000018550
138	HMG3	TRCN0000019105 TRCN0000019104 TRCN0000019108 TRCN0000019106 TRCN0000019107
139	HTATIP/KAT5	TRCN0000020314 TRCN0000020315 TRCN0000020316 TRCN0000020317 TRCN0000020318
140	ING1	TRCN0000073268 TRCN0000073269 TRCN0000073270 TRCN0000073271 TRCN0000073272
141	ING2	TRCN0000019214 TRCN0000019215 TRCN0000019216 TRCN0000019217 TRCN0000019218
142	ING3	TRCN0000073273 TRCN0000073277 TRCN0000073274 TRCN0000073275 TRCN0000073276
143	ING4	TRCN0000128673 TRCN0000129127 TRCN0000129334 TRCN0000129395 TRCN0000131049 TRCN0000103947
144	ING5	TRCN0000020089 TRCN0000020091 TRCN0000020090 TRCN0000020092

		TRCN0000020093
145	INOC1	TRCN0000107555 TRCN0000107556 TRCN0000107557 TRCN0000107558 TRCN0000107559
146	JARID1A	TRCN0000014628 TRCN0000014629 TRCN0000014630 TRCN0000014631 TRCN0000014632
147	JARID1B	TRCN0000014758 TRCN0000014759 TRCN0000014760 TRCN0000014761 TRCN0000014762
148	JARID1C	TRCN0000022084 TRCN0000022085 TRCN0000022084 TRCN0000022085 TRCN0000022087 TRCN0000022088
149	JARID1D	TRCN0000022114 TRCN0000022115 TRCN0000022116 TRCN0000022117 TRCN0000022118
150	JARID2	TRCN0000107365 TRCN0000107366 TRCN0000107367 TRCN0000107368 TRCN0000107369
151	JMJD1A	TRCN0000021149 TRCN0000021150 TRCN0000021151 TRCN0000021152 TRCN0000021153
152	JMJD1B	TRCN0000017093 TRCN0000017094 TRCN0000017095

		TRCN0000017096 TRCN0000017097
153	JMJD1C	TRCN0000107560 TRCN0000107561 TRCN0000107562 TRCN0000107563 TRCN0000107564
154	JMJD2A	TRCN0000013493 TRCN0000013494 TRCN0000013495 TRCN0000013496 TRCN0000013497
155	JMJD2B	TRCN0000018013 TRCN0000018014 TRCN0000018015 TRCN0000018016 TRCN0000018017
156	JMJD2C	TRCN0000022054 TRCN0000022055 TRCN0000022056 TRCN0000022057 TRCN0000022058
157	JMJD2D	TRCN0000149641 TRCN0000149414 TRCN0000147186 TRCN0000148837 TRCN0000148192
158	JMJD3/KDM6B	TRCN0000095268
159	JMJD4	TRCN0000138244 TRCN0000138555 TRCN0000138689 TRCN0000137919 TRCN0000138740
160	JMJD5	TRCN0000129536 TRCN0000128906 TRCN0000130971 TRCN0000130438 TRCN0000130552
161	JMJD6	TRCN0000063338 TRCN0000063339

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162	L3MBTL	TRCN0000016863 TRCN0000016864 TRCN0000016865 TRCN0000016866 TRCN0000016867
163	L3MBTL2	TRCN0000021725 TRCN0000021728 TRCN0000021726 TRCN0000021727 TRCN0000021724
164	MBD1	TRCN0000015428 TRCN0000015429 TRCN0000015430 TRCN0000015431 TRCN0000015432
165	MBD2	TRCN0000013318 TRCN0000013319 TRCN0000013320 TRCN0000013321 TRCN0000013322
166	MBD3	TRCN0000017353 TRCN0000017354 TRCN0000017355 TRCN0000017356 TRCN0000017357
167	MBD4	TRCN0000013313 TRCN0000013314 TRCN0000013315 TRCN0000013316 TRCN0000013317
168	MBTD1	TRCN00000153466 TRCN00000152974 TRCN00000151070 TRCN00000150313 TRCN00000154234 TRCN0000078113 TRCN0000073848

		TRCN0000097657 TRCN0000097655
169	MDC1	TRCN0000018850 TRCN0000018851 TRCN0000018852 TRCN0000018853 TRCN0000018854
170	MDS1	TRCN0000015583 TRCN0000015584 TRCN0000015585 TRCN0000015586 TRCN0000015587
171	MECP2	TRCN0000021239 TRCN0000021240 TRCN0000021241 TRCN0000021242 TRCN0000021243
172	MEN1	TRCN0000040138 TRCN0000040139 TRCN0000040140 TRCN0000040141 TRCN0000040142
173	METTL8	TRCN0000007228
174	MGMT	TRCN0000022364 TRCN0000022365 TRCN0000022366 TRCN0000022367 TRCN0000022368
175	MIER1	TRCN0000108032 TRCN0000108030 TRCN0000108031 TRCN0000108033 TRCN0000108034 TRCN0000095734 TRCN0000095737
176	MIER2	TRCN0000137269 TRCN0000138895 TRCN0000138939 TRCN0000137227 TRCN0000137705

177	MIER3	TRCN0000127638 TRCN0000130779 TRCN0000129622 TRCN0000128394 TRCN0000129860
178	MLL1	TRCN0000108069
179	MLL3	TRCN0000008742 TRCN0000008743 TRCN0000008744 TRCN0000008745 TRCN0000008746 TRCN0000008746
180	MLL4	TRCN0000005958 TRCN0000005959 TRCN0000005960 TRCN0000005961 TRCN0000005962
181	MLL4/WBP7	TRCN0000005958 TRCN0000005959 TRCN0000005960 TRCN0000005961 TRCN0000005962
182	MLL5	TRCN0000150550 TRCN0000154711 TRCN0000155824 TRCN0000157609 TRCN0000158279
183	MLLT6	TRCN0000019902 TRCN0000019899 TRCN0000019901 TRCN0000019903 TRCN0000019900 TRCN0000074123
184	MORF4L1	TRCN0000107580 TRCN0000107581 TRCN0000107582 TRCN0000107583 TRCN0000107584 TRCN0000015007
185	MORF4L2	TRCN0000107309



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186	MPHOSPH8	TRCN0000117922 TRCN0000117925 TRCN0000117923 TRCN0000117926 TRCN0000117924 TRCN0000085440
187	MSL3L1/MSL3	TRCN0000022104 TRCN0000022105 TRCN0000022106 TRCN0000022107 TRCN0000022108
188	MTA1	TRCN0000013358 TRCN0000013359 TRCN0000013360 TRCN0000013361 TRCN0000013362
189	MTA2	TRCN0000013373 TRCN0000013374 TRCN0000013375 TRCN0000013376 TRCN0000013377
190	MTA3	TRCN0000016458 TRCN0000016459 TRCN0000016460 TRCN0000016461 TRCN0000016462
191	MTF2	TRCN0000107680 TRCN0000107681 TRCN0000107682 TRCN0000107683 TRCN0000107684 TRCN0000086133 TRCN0000086137
192	MYST1	TRCN0000034875 TRCN0000034876

		TRCN0000034877 TRCN0000034878
193	MYST2	TRCN0000021629 TRCN0000021630 TRCN0000021631 TRCN0000021632 TRCN0000021633
194	MYST3	TRCN0000013123 TRCN0000013124 TRCN0000013125 TRCN0000013126
195	MYST4	TRCN0000019359 TRCN0000019360 TRCN0000019361 TRCN0000019362 TRCN0000019363
196	N6AMT1	TRCN0000038755 TRCN0000038758 TRCN0000038754 TRCN0000038757 TRCN0000038756
197	N6AMT2	TRCN0000139966 TRCN0000122732 TRCN0000140564 TRCN0000122562 TRCN0000145142
198	NAP1I1	TRCN0000149610 TRCN0000149872 TRCN0000148741 TRCN0000149921 TRCN0000155836 TRCN0000092890
199	NAP1L2	TRCN0000147762 TRCN0000146531 TRCN0000146873 TRCN0000148173 TRCN0000147038
200	NAP1L3	TRCN0000149156 TRCN0000147100 TRCN0000149498

		TRCN0000147275 TRCN0000149864
201	NAP1L4	TRCN0000151475 TRCN0000154499 TRCN0000155016 TRCN0000151026 TRCN0000156041
202	NCOA1	TRCN0000004082 TRCN0000004080 TRCN0000004078 TRCN0000004079 TRCN0000095569 TRCN0000004081
203	NCOA3	TRCN0000019699 TRCN0000019700 TRCN0000019701 TRCN0000019702 TRCN0000019703
204	NCOA6	TRCN0000063267 TRCN0000063264 TRCN0000063266 TRCN0000063263 TRCN0000063265 TRCN0000176319
205	NCOR1	TRCN0000060654 TRCN0000060657 TRCN0000060656 TRCN0000060653 TRCN0000060655
206	NCOR2	TRCN0000060704 TRCN0000060706 TRCN0000060703 TRCN0000060707 TRCN0000060705 TRCN0000063970 TRCN0000095279
207	NROB2	TRCN0000033649 TRCN0000033650 TRCN0000033651 TRCN0000033652

		TRCN0000033653
208	NSD1	TRCN0000061353 TRCN0000061354 TRCN0000061356 TRCN0000061357
209	PADI4	TRCN0000051678 TRCN0000051679 TRCN0000051680 TRCN0000051681
210	PARP1	TRCN0000007928 TRCN0000007929 TRCN0000007930 TRCN0000007931 TRCN0000007932
211	PARP2	TRCN0000007933 TRCN0000007934 TRCN0000007935 TRCN0000007936 TRCN0000007937
212	PAX5	TRCN0000016058 TRCN0000016059 TRCN0000016060 TRCN0000016061 TRCN0000016062
213	PAXIP1	TRCN0000128479 TRCN0000150101 TRCN0000147957 TRCN0000147635 TRCN0000129174
214	PBRM1	TRCN0000015995 TRCN0000015994 TRCN0000015993 TRCN0000015997 TRCN0000007228
215	PCAF/KAT2B	TRCN0000018528 TRCN0000018529 TRCN0000018530 TRCN0000018531 TRCN0000018532
216	PCGF1	TRCN0000073131

		TRCN0000073128 TRCN0000073129 TRCN0000073130 TRCN0000073132
217	PCGF2	TRCN0000022029 TRCN0000022030 TRCN0000022031 TRCN0000022032 TRCN0000022033
218	PCGF3	TRCN0000095494
219	PCGF5	TRCN0000073113 TRCN0000073115 TRCN0000073117 TRCN0000073116 TRCN0000140719 TRCN0000095162 TRCN0000073114
220	PCGF6	TRCN0000073108 TRCN0000073109 TRCN0000073112 TRCN0000073110 TRCN0000073111 TRCN0000095301
221	PCMT1	TRCN0000036399 TRCN0000036400 TRCN0000036401 TRCN0000036402 TRCN0000036403
222	PCNA	TRCN0000003861 TRCN0000003862 TRCN0000003863 TRCN0000003864 TRCN0000010826
223	PHF1	TRCN0000019184 TRCN0000019185 TRCN0000019186 TRCN0000019187 TRCN0000019188
224	PHF11	TRCN0000020112 TRCN0000020110

		TRCN0000020111 TRCN0000020113 TRCN0000020109 TRCN0000073848
225	PHF12	TRCN0000015705 TRCN0000015706 TRCN0000015707 TRCN0000015704 TRCN0000015703 TRCN0000007228 TRCN0000084422 TRCN0000084419
226	PHF13	TRCN0000019637 TRCN0000019635 TRCN0000019634 TRCN0000019636 TRCN0000019638
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230	PHF17	TRCN0000019621 TRCN0000019619 TRCN0000019620 TRCN0000019622 TRCN0000019623
231	PHF19	TRCN00000108185 TRCN00000108188 TRCN00000108189

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232	PHF20	TRCN0000016316 TRCN0000016313 TRCN0000016317 TRCN0000016314 TRCN0000016315
233	PHF21A	TRCN0000037003 TRCN0000037000 TRCN0000036999 TRCN0000037001 TRCN0000037002 TRCN0000124326
234	PHF23	TRCN0000019259 TRCN0000019263 TRCN0000019261 TRCN0000019262 TRCN0000019260
235	PHF3	TRCN0000019114 TRCN0000019115 TRCN0000019116 TRCN0000019117 TRCN0000019118
236	PHF8	TRCN0000118317 TRCN0000118318 TRCN0000118319 TRCN0000118320 TRCN0000118321
237	PML	TRCN0000003866 TRCN0000003867 TRCN0000003868 TRCN0000003869
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239	PPARGC1A	TRCN0000001165 TRCN0000001166 TRCN0000001167

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240	PRDM1	TRCN0000013608 TRCN0000013609 TRCN0000013610 TRCN0000013611 TRCN0000013612
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250	PRDM8	TRCN0000016758 TRCN0000016759 TRCN0000016760 TRCN0000016761 TRCN0000016762
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253	PRKAA2	TRCN0000002168 TRCN0000002169 TRCN0000002170 TRCN0000002171 TRCN0000002172
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256	PRMT2	TRCN0000035894 TRCN0000035895 TRCN0000035896 TRCN0000035897 TRCN0000035898
257	PRMT3	TRCN0000034569 TRCN0000034570 TRCN0000034571 TRCN0000034572 TRCN0000034573
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260	PRMT7	TRCN0000035149 TRCN0000035150 TRCN0000035151 TRCN0000035152 TRCN0000035153
261	PRMT8	TRCN0000034749 TRCN0000034750 TRCN0000034751 TRCN0000034752 TRCN0000034753
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265	RAG2	TRCN0000051228 TRCN0000051229 TRCN0000051230 TRCN0000051231 TRCN0000051232
266	RBBP4	TRCN0000115870 TRCN0000115867 RCN0000115868 TRCN0000115869 TRCN0000115871
267	RBBP5	TRCN0000034410 TRCN0000034413
268	RBBP7	TRCN0000038888 TRCN0000038887 TRCN0000038884 TRCN0000038886 TRCN0000038885
269	RCOR1	TRCN0000128570 TRCN0000129660 TRCN0000147184 TRCN0000128260 TRCN0000147902
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272	RERE	TRCN0000013578 TRCN0000013579

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273	RING1	TRCN0000021989 TRCN0000021990 TRCN0000021991 TRCN0000021992 TRCN0000021993
274	RNF2	TRCN0000033694 TRCN0000033695 TRCN0000033696 TRCN0000033697 TRCN0000033698
275	RNF20	TRCN0000033874 TRCN0000033875 TRCN0000033876 TRCN0000033877 TRCN0000033878
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278	RPA1	TRCN0000010985 TRCN0000005984 TRCN0000010982 TRCN0000010983 TRCN0000005985
279	RPA3	TRCN0000018860 TRCN0000018861 TRCN0000018862 TRCN0000018863 TRCN0000018864
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287	SETD2	TRCN0000003029 TRCN0000003030 TRCN0000003031 TRCN0000003032 TRCN0000003033
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289	SETD5	TRCN0000098703 TRCN0000098704
290	SETD7	TRCN0000078628 TRCN0000078629 TRCN0000078630 TRCN0000078631 TRCN0000078632
291	SETD8	TRCN0000130139 TRCN0000130036 TRCN0000148268 TRCN0000128082
292	SFMBT1	TRCN0000146760 TRCN0000146761 TRCN0000147760 TRCN0000148746 TRCN0000147815
293	SFMBT2	TRCN0000148663 TRCN0000149533 TRCN0000149620 TRCN0000149672 TRCN0000146511
294	SIN3A	TRCN0000021774 TRCN0000021775 TRCN0000021776 TRCN0000021777 TRCN0000021778
295	SIN3B	TRCN0000107807 TRCN0000107805 TRCN0000107809 TRCN0000107808 TRCN0000107806

296	SIRT1	TRCN0000018979 TRCN0000018980 TRCN0000018981 TRCN0000018982 TRCN0000018983
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308	SMARCAL1	TRCN0000083572 TRCN0000083568 TRCN0000083569 TRCN0000083570 TRCN0000083571
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310	SMARCC1	TRCN0000015628 TRCN0000015631 TRCN0000015629 TRCN0000015630 TRCN0000015632
311	SMARCC2	TRCN0000015698 TRCN0000015699 TRCN0000015701 TRCN0000015702



312	SMARCD1	TRCN0000151662 TRCN0000157022 TRCN0000153623 TRCN0000158153 TRCN0000157443 TRCN0000092957
313	SMARCD2	TRCN0000021267 TRCN0000021266 TRCN0000021265 TRCN0000021264 TRCN0000021268
314	SMARCE1	TRCN0000015778 TRCN0000015779 TRCN0000015780 TRCN0000015781 TRCN0000015782
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317	SMYD3	TRCN0000123289 TRCN0000123292 TRCN0000123293 TRCN0000123290 TRCN0000123291
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320	SUDS3	TRCN0000038912 TRCN0000038909 TRCN0000038910 TRCN0000038911 TRCN0000038913 TRCN0000039314
321	SUPT16H	TRCN0000001257 TRCN0000001258 TRCN0000001259 TRCN0000001260 TRCN0000001261
322	SUV39H1	TRCN0000158337 TRCN0000150622 TRCN0000158270 TRCN0000157251 TRCN0000157285
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325	Suv420h2	TRCN0000141493 TRCN0000141926 TRCN0000143766 TRCN0000142036 TRCN0000141258 TRCN0000039200
326	SUZ12	TRCN0000038724 TRCN0000038725 TRCN0000038726 TRCN0000038727 TRCN0000038728
327	TADA2L/TADA2A	TRCN0000017299 TRCN0000017298

		TRCN0000017302 TRCN0000017301 TRCN0000017300
328	TADA3/TADA3L	TRCN0000015737 TRCN0000015733 TRCN0000015736 TRCN0000015734 TRCN0000015735
329	TAF1	TRCN0000006284 TRCN0000006285 TRCN0000006286 TRCN0000006287 TRCN0000006288
330	TAF3	TRCN0000016608 TRCN0000016609 TRCN0000016610 TRCN0000016611 TRCN0000016612
331	TBL1X	TRCN0000118647 TRCN0000118648 TRCN0000118649 TRCN0000118650 TRCN0000118651
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333	TET1	TRCN0000075027 TRCN0000075025 TRCN0000075024 TRCN0000075026 TRCN0000075023
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335	TLK1	TRCN0000007056 TRCN0000007057

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337	TRIM24	TRCN0000021260 TRCN0000021259 TRCN0000021262 TRCN0000021263 TRCN0000021261 TRCN0000088522
338	TRIM28	TRCN0000017999 TRCN0000018002 TRCN0000018000 TRCN0000018001 TRCN0000017998
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340	TRP53BP1	TRCN0000081779
341	TRRAP	TRCN0000005363 TRCN0000005361 TRCN0000005362 TRCN0000005364 TRCN0000005365
342	UBE2A	TRCN0000010842 TRCN0000004005 TRCN0000004006 TRCN0000004007 TRCN0000004008
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345	UBE2I	TRCN0000011077 TRCN0000007205 TRCN0000007206 TRCN0000011078 TRCN0000007207 TRCN0000011078 TRCN0000011077
346	UHRF1	TRCN0000004353 TRCN0000004352 TRCN0000004356 TRCN0000004354 TRCN0000004355 TRCN00000112188
347	UHRF1BP1	TRCN0000064774 TRCN0000064777 TRCN0000064776 TRCN0000064773 TRCN0000064775 TRCN0000073848
348	UHRF2	TRCN0000003479 TRCN0000003481 TRCN0000003480 TRCN0000003482 TRCN0000010793 TRCN0000040626
349	USP22	TRCN0000046924 TRCN0000046925 TRCN0000046926 TRCN0000046927
350	USP51	TRCN0000038849 TRCN0000038850 TRCN0000038851 TRCN0000038852 TRCN0000038853

351	UTX	TRCN0000107760 TRCN0000107761 TRCN0000107762 TRCN0000107763 TRCN0000107764
352	WDR5	TRCN0000118047 TRCN0000118048 TRCN0000118049 TRCN0000118050 TRCN0000118051
353	WDR54	TRCN0000072663 TRCN0000072664 TRCN0000072665 TRCN0000072666 TRCN0000072667
354	WDR82	TRCN0000130689 TRCN0000129877 TRCN0000128304 TRCN0000127796 TRCN0000130791
355	WHSC1	TRCN0000019814 TRCN0000019815 TRCN0000019816 TRCN0000019817 TRCN0000019818
356	WHSC1L1	TRCN0000015613 TRCN0000015614 TRCN0000015615 TRCN0000015616 TRCN0000015617
357	WNT5A	TRCN0000062713 TRCN0000062714 TRCN0000062715 TRCN0000062716 TRCN0000062717
358	YEATS4	TRCN0000013143 TRCN0000013144 TRCN0000013146 TRCN0000013145 TRCN0000013147

359	ZCWPW1	TRCN0000139148 TRCN0000140908 TRCN0000145000 TRCN0000139000 TRCN0000140405
360	ZFPM1	TRCN0000107406 TRCN0000107408 TRCN0000107407 TRCN0000107405 TRCN0000107409
361	ZMYND11	TRCN0000021253 TRCN0000021251 TRCN0000021252 TRCN0000021249 TRCN0000021250 TRCN0000088135
362	ZMYND8	TRCN0000037997 TRCN0000037996 TRCN0000037998 TRCN0000037995 TRCN0000037994 TRCN0000007228
363	ZNFX1	TRCN0000017371 TRCN0000017369 TRCN0000017372 TRCN0000017368 TRCN0000017370 TRCN0000073848

**Table S2:** List of genes for which multiple shRNAs were isolated from the epigenome-wide shRNA screen.

<b>S.No.</b>	<b>Gene Symbol</b>	<b>Gene Name</b>	<b>Accession Number</b>
<b>1</b>	<b><i>BOP1</i></b>	Block of Proliferation	NM_015201.4
<b>2</b>	<b><i>HAT1</i></b>	Histone acetyltransferase 1	NM_003642.3
<b>3</b>	<b><i>HMGB1</i></b>	High mobility group protein B1	NM_001313892
<b>4</b>	<b><i>HMGNI</i></b>	High mobility group nucleosomal binding protein 1	NM_004965.6
<b>5</b>	<b><i>ING5</i></b>	Inhibitor of growth 5	NM_001330161.1
<b>6</b>	<b><i>KDM4C</i></b>	Lysine-specific demethylase 4C	NM_001146695



**Table S3:** Patient sample information used for BOP1 expression analysis.

S.No.	SAMPLE ID	STAGE	MUTATION	TREATMENT	RESPONSE	SAMPLES ANALYZED	BOP1 PROTEIN EXPRESSION IN PROGRESSED SAMPLES
Pair 1	#1218	IV M1c	V600E	Seq. BRAFi + MEKi	SD	Matched Pre- and progressed	Downregulated
Pair 2	#1117	IV M1c	V600E	Dual BRAFi + MEKi	PR	Matched Pre- and progressed	Downregulated
Pair 3	#848	IV M1c	V600E	Dual BRAFi + MEKi	PR	Matched Pre- and progressed	Downregulated
Pair 4	#757	IV M1c	V600E	BRAFi	PR	Matched Pre- and progressed	Downregulated
Pair 5	#0470	IV M1c	V600K	BRAFi	PR	Matched Pre- and progressed	Downregulated
Pair 6	#0959	IV M1a	V600E	BRAFi	PR	Matched Pre- and progressed	Downregulated
Pair 7	#0923	IV M1c	V600E	Dual BRAFi + MEKi	PR	Matched Pre- and progressed	Downregulated
Pair 8	#2132	IV M1c	V600E	Dual BRAFi + MEKi	PR	Matched Pre- and progressed	Unchanged
Pair 9	#1351	IV M1a	V600E	Seq. BRAFi + MEKi	PR	Matched Pre- and progressed	Unchanged
Pair 10	#1577	IV M1c	V600E	Dual BRAFi + MEKi	PR	Matched Pre- and progressed	Upregulated
Pair 11	#1254	IV M1c	V600E	BRAFi	PR	Matched Pre- and progressed	Upregulated

SD: Stable disease  
PR: Partial response  
Seq: Sequential

**Table S4:** Primer sequences, clone IDs, catalog numbers, antibodies, and chemical inhibitors used in this study. Primers were used for qRT-PCR analysis. The shRNAs used herein were obtained from Open Biosystems; clone IDs and catalog numbers are listed. The antibodies were used for immunoblot analyses. The source and concentrations of chemical inhibitors used for drug treatment experiments are summarized.

Application	Gene symbol	Forward primer (5'-3')	Reverse primer (5'-3')
RT-qPCR	<i>BOP1</i>	aggactcggctgctgctg	cccacttggcgctcctcctc
	<i>HAT1</i>	tggcagatagaggcacaacag	acacgccggaatctccac
	<i>KDM4C</i>	gcagacaccgggtaagagg	cttctcgcaagcctggact
	<i>HMGB1</i>	ggagaacatcctggcctgtc	cagcttcgcagcctctttt
	<i>HMGN1</i>	aaacaggccgaagtggctaa	agggagacaggaccactga
	<i>ING5</i>	cccaaacgaaccacgtact	gggttctggtgaaggcca
	<i>DUSP1</i>	tgtggaggacaaccacaagg	aaactcaaaggcctctgcca
	<i>DUSP2</i>	ctgtcccgatctgtgctctg	ccatctggtgtcctccaca
	<i>DUSP4</i>	gaggggaatggacaccacat	tgtctcctccaaagccatt
	<i>DUSP5</i>	ggatccctgtggaagacagc	gaccatgctcctctgct
	<i>DUSP6</i>	ttgagacgctcgtgtttgt	agccgctggctcttagtgc
	<i>DUSP7</i>	ctcgcaagtatggcatcaa	ggtcagagatggggatctgc
	<i>DUSP8</i>	cgccctctctgtgttctt	gacagaggaacgggaacag
	<i>DUSP9</i>	cacacatctgtgggctgtt	cacaggggtgtggacagaaa
	<i>DUSP10</i>	ccgacctcaggatctcaacc	cctgtcgtaggtgcccaca
	<i>DUSP16</i>	cagactccgaaccagctcc	ccagcagactctgtgagggtg
	<i>MX-YTX</i>	caccaagagggtggaagagg	ctccaagggcagaaggaatg
	<i>SPRY1</i>	ccaagcaactgatttgga	Gtagggctctgggagcagtg
	<i>SPRY2</i>	agtcgtctccagctccgaac	Cctgagctctggcctccat
	<i>PP2A</i>	atcttgcctgcatcaagga	Tctttgccagaatgggaga
	<i>PP2C</i>	tgtcatgcacatctgtctg	tcagcagggttcttctcc
	<i>HePTP</i>	caccacacttggccctgta	acttcccagaccacctt
	<i>DUSP3</i>	gtgtgatggatgtgtggg	tcattgtcctggctgagt
	<i>PTP-SL</i>	ggcataaagggtgctctgc	agcctcgaatgggacaga
	<i>STEP</i>	tgtgtcctcagccaggaa	tgccctaactcccgtcatct
	<i>CIRH1A</i>	caggcagcgtgtcataag	gtcccagaactgcacctcc
	<i>DKC1</i>	tgggagttggtgctcagatg	tcaatgccgtcctcatalcg
	<i>EMG1</i>	accgaattccagaacctt	ggccccctaccacaaaaacaa
	<i>HCA66</i>	aagggtccaaaagccaagaa	ctgaaaatggtcggctctcc
	<i>NHP2</i>	tcacgcggaagctctacaaa	gagccccctatagggtagg
	<i>RBM28</i>	cctaccaaagccagcgttc	gtccggccttttcagtcac
	<i>RMRP</i>	acgtagacattcccgcctc	acgcactgcctcgtaacta
<i>RPL11</i>	caggttcagcatcgagac	cgggaaattattgcccagga	
<i>RPL35A</i>	aaggccattttgtggcta	gaaccatgccactgttcca	
<i>RPL36</i>	caacaagggccacaaagtga	gggaggggctcagctttct	
<i>RPL5</i>	cgaaacgcttggatgataca	gtcacctccacttggccttc	
<i>RPS14</i>	tgtgactggtggatgaagg	ggtgataccagctcctgctc	
<i>RPS15</i>	agaagccggaagtgtgaag	atggagaactgcccaggta	

	<i>RPS17</i>	gccattatcccagcaaaaa	gccattatcccagcaaaaa
	<i>RPS19</i>	acgaacttcgccctgagag	gggagcaagctctttgtgct
	<i>RPS24</i>	tttggtggtggcaagacaac	ttgcagtccccctgacttt
	<i>RPS27A</i>	cattgggtggtgcgctaa	ctcaggggtgatggcttcc
	<i>RPS7</i>	tcccaggagaaagccatggt	ctccagaagagcctgggaga
	<i>SBDS</i>	gacgctgctgaagctagt	taccacggccacattggta
	<i>TCOF1</i>	tgagctgctggaacaggaaa	ccttgaagtcgtggaggtc
	<i>UTP14C</i>	aaagcccagtgccaagaaaa	aggggtctggatggctctt
	<i>WDR36</i>	caaggaagcctgcagtgtg	aatatggccacatgggcttc
	<i>NOP10</i>	ctcgttctcccagatgac	agggtcacagtcggaagag
	<i>WDR12</i>	tggctcagctcacaacacgct	ggcaattcagaggcagcagggga
	<i>PES1</i>	gggcacctacgcgttgact	tccacctcggcctcctctc
	<i>28S</i>	agaggtaaacgggtgggtc	ggggcgggagggaacgg
	<i>U1</i>	ccatgatcacgaaggtggtt	atgcagtcgagttcccacat
	<i>SP6</i>	catacgatttagtgacactatag	
	<i>MF18</i>	tacgatacaaggctgttagagag	
	<i>MF19</i>	cgaaccgcaaggaaccttc	
<b>shRNAs</b>	<b>Gene symbol</b>	<b>Clone ID</b>	<b>Catalog number</b>
	<i>BOP1</i>	TRCN0000078069	RHS3979-97066366
		TRCN0000078071	RHS3979-97066368
	<i>HAT1</i>	TRCN0000034735	RHS3979-9602143
		TRCN0000034737	RHS3979-9602145
	<i>KDM4C</i>	TRCN0000022056	RHS3979-9589464
		TRCN0000022058	RHS3979-9589466
	<i>HMGB1</i>	TRCN0000018932	RHS3979-9586254
		TRCN0000018934	RHS3979-9586256
	<i>HMG1</i>	TRCN0000019059	RHS3979-9586467
		TRCN0000019060	RHS3979-9586468
	<i>ING5</i>	TRCN0000020090	RHS3979-9587498
		TRCN0000020092	RHS3979-9587500
	<i>DUSP4</i>	TRCN0000006842	RHS3979-9575763
		TRCN0000006843	RHS3979-9575764
	<i>DUSP6</i>	TRCN0000002437	RHS3979-9571003
		TRCN0000010706	RHS3979-9571004
	<i>RPL11</i>	TRCN0000117714	RHS3979-98059637
		TRCN0000117716	RHS3979-98059653
	<i>RPS24</i>	TRCN0000117547	RHS3979-98059466
		TRCN0000117548	RHS3979-98059474
	<i>WDR12</i>	V3LHS_311782	RHS4430-101164454
		V2LHS_156548	RHS4430-99613637
	<i>PES1</i>	V3LHS_304461	RHS4430-101034784
		V3LHS_304460	RHS4430-101031539

	<b>Protein symbol</b>	<b>Antibody source</b>	<b>Dilution</b>
<b>Immunoblotting</b>	Actin	Cell signaling	1:5000
	KDM4C	Active Motif	1:1000
	HMGB1	Cell signaling	1:2000
	ING5	Santa Cruz Biotechnology	1:200
	HAT1	Santa Cruz Biotechnology	1:200
	P-ERK1/2	Cell signaling	1:2000
	T-ERK1/2	Cell signaling	1:2000
	DUSP4	Cell signaling	1:2000
	BOP1	Santa Cruz Biotechnology	1:200
	V5	Cell signaling	1:2000
	RPL11	Cell signaling	1:2000
	RPS24	Thermofischer scientific	1:1000
	TP53	Santa Cruz Biotechnology	1:200
	HMG1	Active Motif	1:1000
<b>Immuno-fluorescence</b>	GM113	BD Biosciences	1:200
	BOP1	Sigma Aldrich	1:250
	S100	Biogenex Laboratories	1:100
	HMB45	Biogenex Laboratories	1:100
	Alexaflour546	Invitrogen	1:100
	Cy5-Thyramide	Perkin Elmer	1:50
<b>Inhibitors</b>		<b>Concentration</b>	<b>Source</b>
	Vemurafenib	0.1 $\mu$ M, 1 $\mu$ M and 2 $\mu$ M	Selleckchem
	Dabrafenib	50 nM and 100 nM	Selleckchem
<b>cDNA/ORF</b>	<b>Accession number</b>	<b>Clone ID</b>	<b>Source</b>
DUSP4	BC002671	ccsbBroad304_06130	Dharmacon
DUSP6	BC037236	ccsbBroad304_00468	Dharmacon