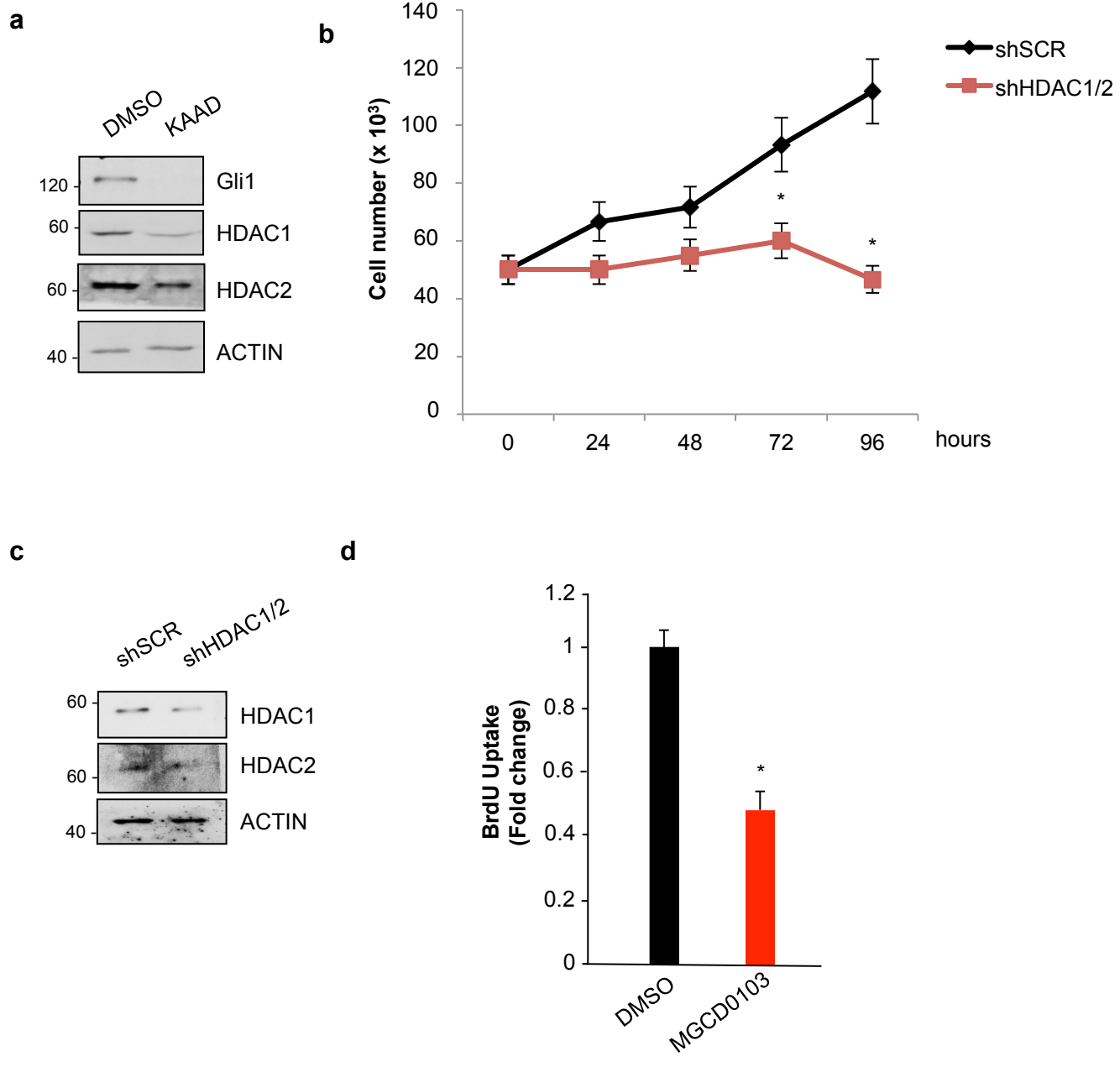


Selective targeting of HDAC1/2 elicits anticancer effects through Gli1 acetylation in preclinical models of SHH Medulloblastoma

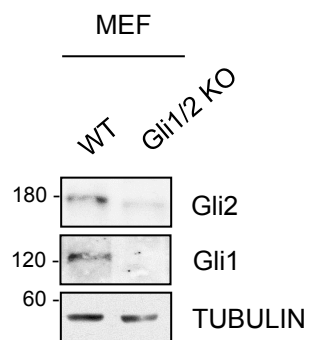
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Supplementary figures, tables and figure legends

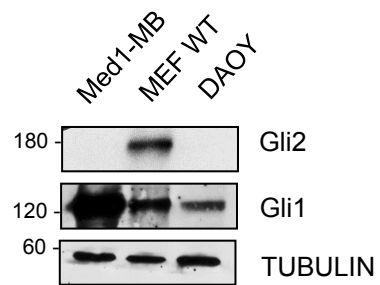


Supplementary Figure S1

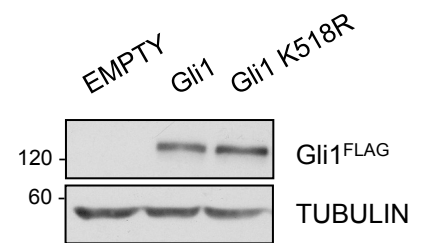
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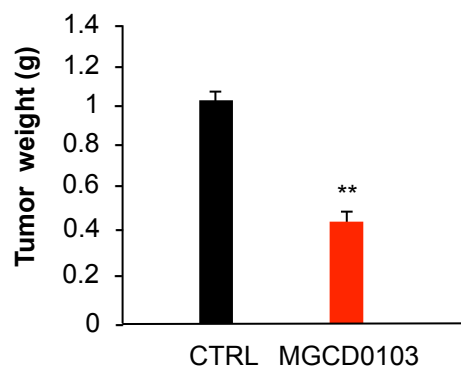
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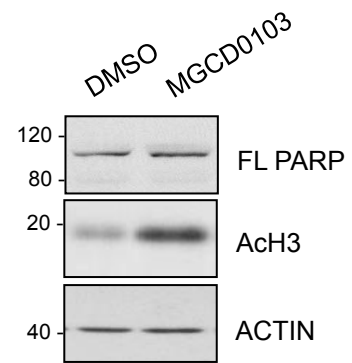
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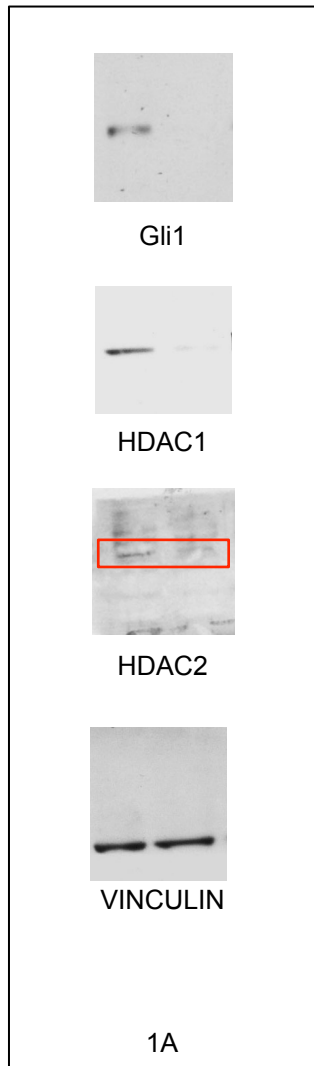
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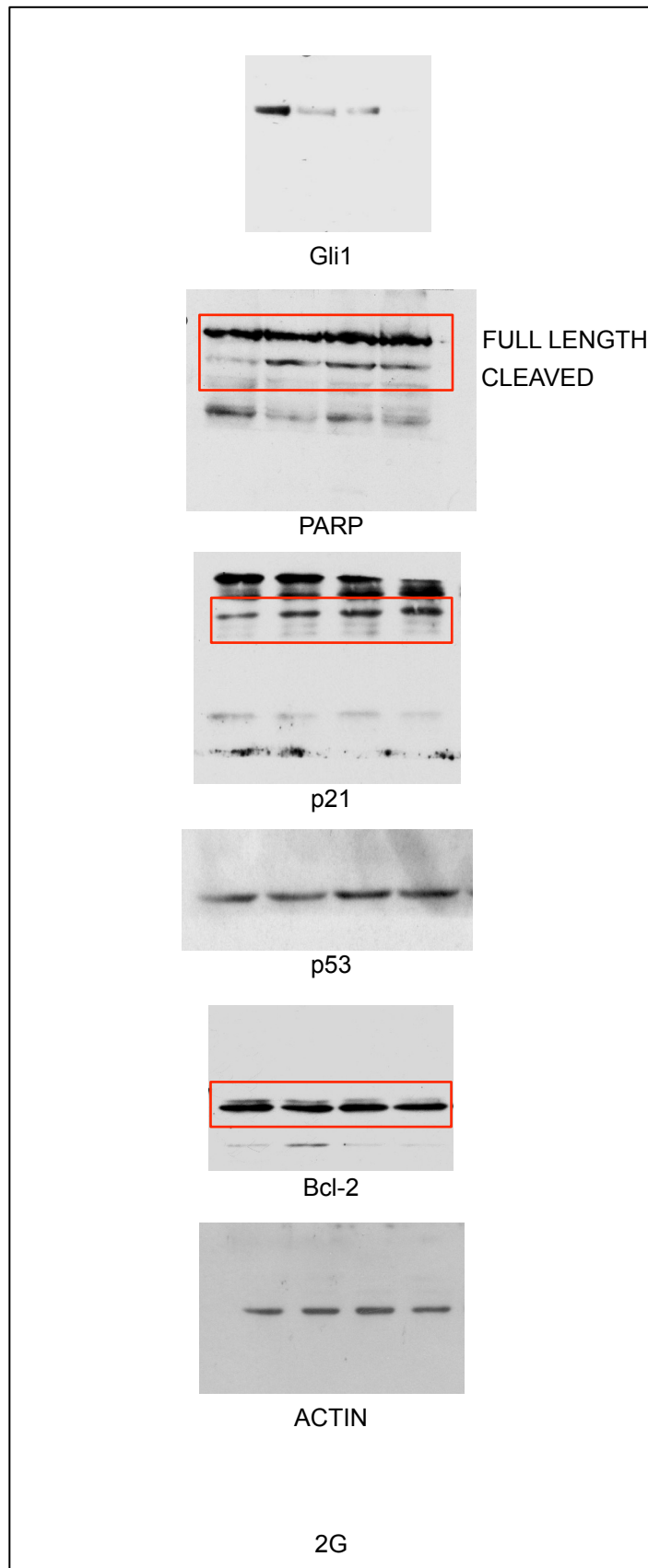
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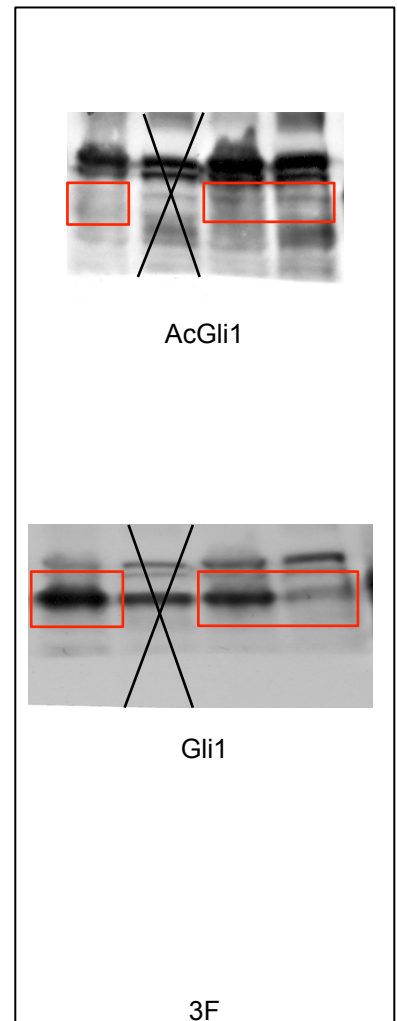
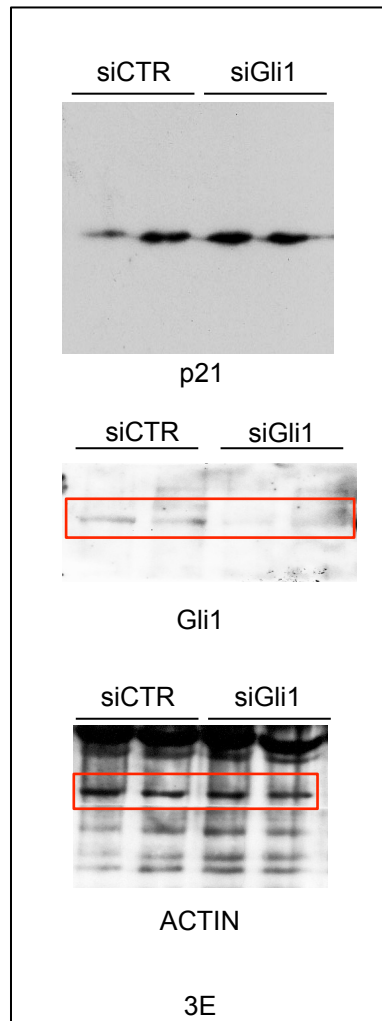
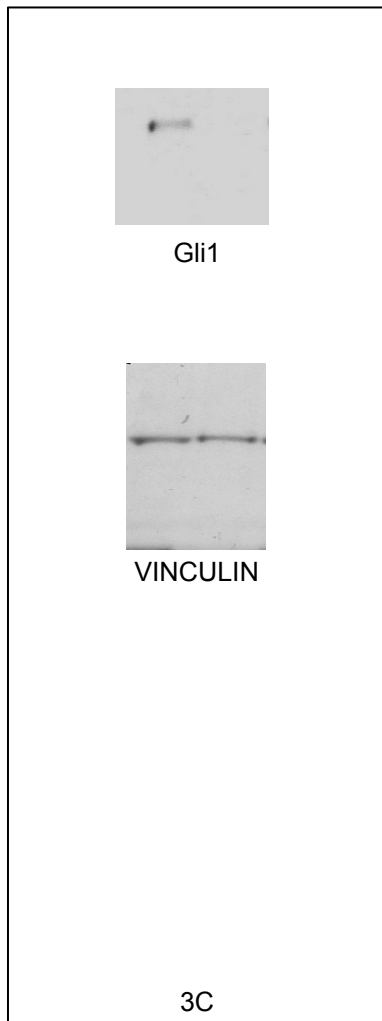
Supplementary Figure S3



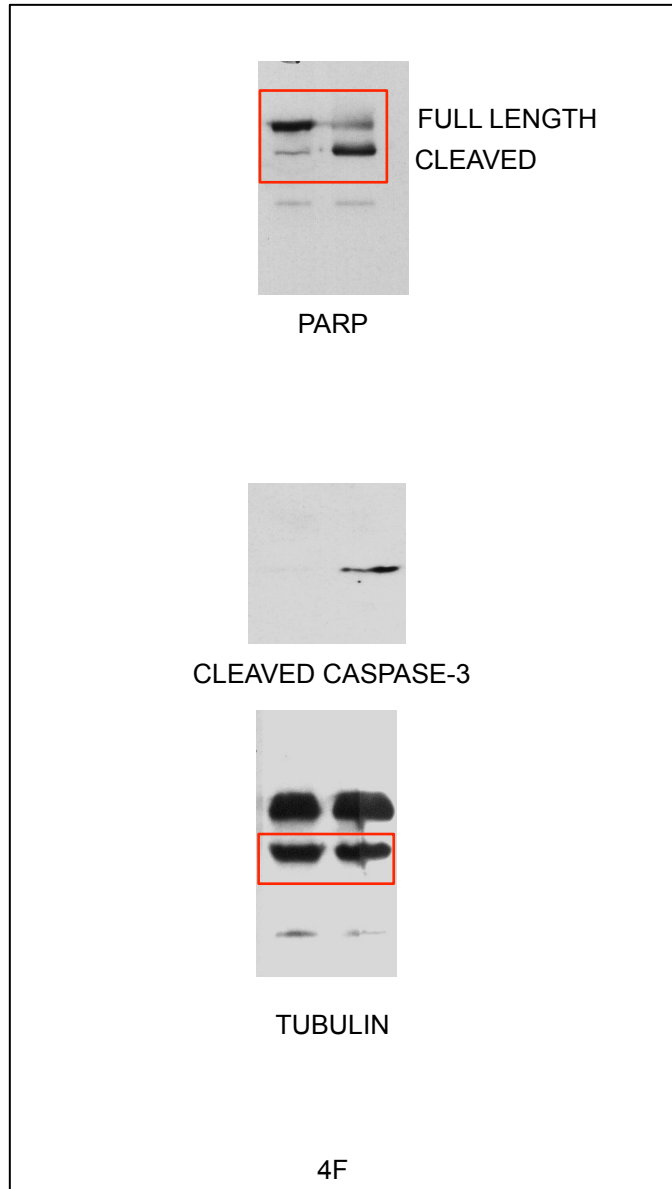
Supplementary Figure S4



Supplementary Figure S5



Supplementary Figure S6



Supplementary Figure S7

Sybr Green oligonucleotides	Sequence
mPtch1	FW: GCATTGGCAGGAGGAGTTGA RW: AGTCATTA ACTGGAACATGGTTTGC
mPtch2	FW: CAACATGGTCGCCTTTTTCA RW: ACCACTATGGCTGCCTGCA
mCyclinD1	FW: TCCGCAAGCATGCACAGA RW: AGGGTGGGTTGGAAATGAACT
mCyclinD2	FW: TTTCAAGTGCCTGCAGAAGG RW: CCAGCCAAGAAACGGTCCAG
mGli1	FW: AAGCCAAC TTTATGTCAGGG RW: AGAGCCCGCTTCTTTCTTAA
mL32	FW: AGAGGTGCTGGGAGCTGCTA RW: GATGGATGGTCTCTGGACGG
mHDAC1	FW: TCGCTGCTGGACTTACGAAA RW: TGTAGGGCAGCTCATTAGGGA
mHDAC2	FW: TCCGGTGTTTGATGGACTCTT RW: TCACAGCCCCAGCAACTGA
mN-Myc	FW: AAGCGTGGATTATGACAGGAAGTC RW: CTTGTCAACAACGCCAGTTTCCAC

Supplementary Table 1

Figure S1: HDAC1 and HDAC2 inhibition impairs Mb primary tumors growth *in vitro*.

(a) Western Blot analysis of Gli1, HDAC1, HDAC2 protein levels in primary MB cells treated with 1 μ M Kaad-cyclopamine or DMSO for 72 hours. Actin, loading control. (b) Growth curve of primary SHH MB cells stably transduced with lentiviruses expressing HDAC1/2 or scrambled shRNAs. Experiments were performed in triplicate for the indicated times. (c) Western blot analysis to verify knockdown efficacy. (d) BrdU incorporation assay in primary MB cells treated for 48 hours with 0.5 μ M MGCD0103 or DMSO. *p<0,05.

Figure S2: Gli levels in cell lines used. (a) Gli1 and Gli2 protein levels on MEF cells WT or Gli1/ Gli2 KO. Tubulin, loading control. (b) Western Blot analysis of Gli1 and Gli2 protein levels in Med1-MB, MEF and DAOY cells. Tubulin, loading control. (c) Western blot analysis in DAOY cells stably expressing Flag-tagged Gli1 WT and K518R (figure 3h). Tubulin, loading control.

Figure S3: Mocetinostat effect in mouse models of SHH MB. (a) Average tumor weight of tumor masses explanted at the end of the allograft experiment shown in figure 4a. (b) Western blot analysis on cerebella from WT littermates from figure 4f. Mice were subcutaneously injected with DMSO or 170mg/Kg MGCD0103 for 6 hours (n= 3 CTRL; n=3 MGCD0103). PARP, AcH3 and actin levels are shown.

Figure S4: Uncropped Western blots related to main Fig.1.

Figure S5: Uncropped Western blots related to main Fig.2.

Figure S6: Uncropped Western blots related to main Fig.3.

Figure S7: Uncropped Western blots related to main Fig.4.

Supplementary table 1. Sybr Green oligonucleotides designed and used in the experiments.