

# Supporting Information

## Discovery of a Small-Molecule Degradator of Bromodomain and Extra-Terminal (BET) Proteins with Picomolar Cellular Potencies and Capable of Achieving Tumor Regression

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**Figure S1.** Western blotting analysis of BRD2, BRD3 and BRD4 proteins in RS4;11 cells treated with compounds **8**, **23** and **26**.

**Figure S2.** <sup>1</sup>H NMR spectrum for compound **23**.

**Figure S3.** <sup>13</sup>C NMR spectrum for compound **23**.

**Figure S4.** UPLC-MS results for compound **23**.

**Figure S5.** Metabolism of compound **23** in mouse liver microsomes.

**Figure S1.** Western blotting analysis of BRD2, BRD3 and BRD4 proteins in RS4;11 cells treated with BET degraders **23** and **26** and BET inhibitor **8**. RS4;11 cells were treated for 3 h with individual compound at indicated concentrations and proteins were probed by specific antibodies. GAPDH was used as the loading control.

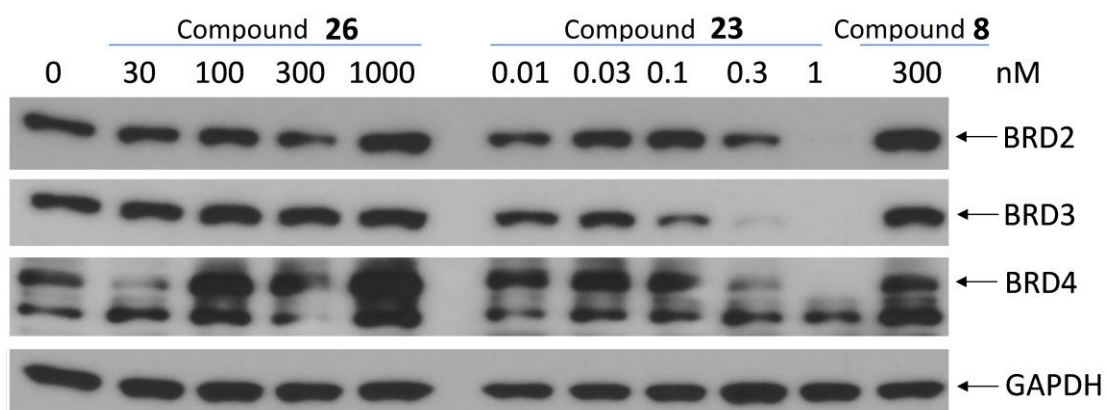


Figure S2. <sup>1</sup>H NMR of 23.

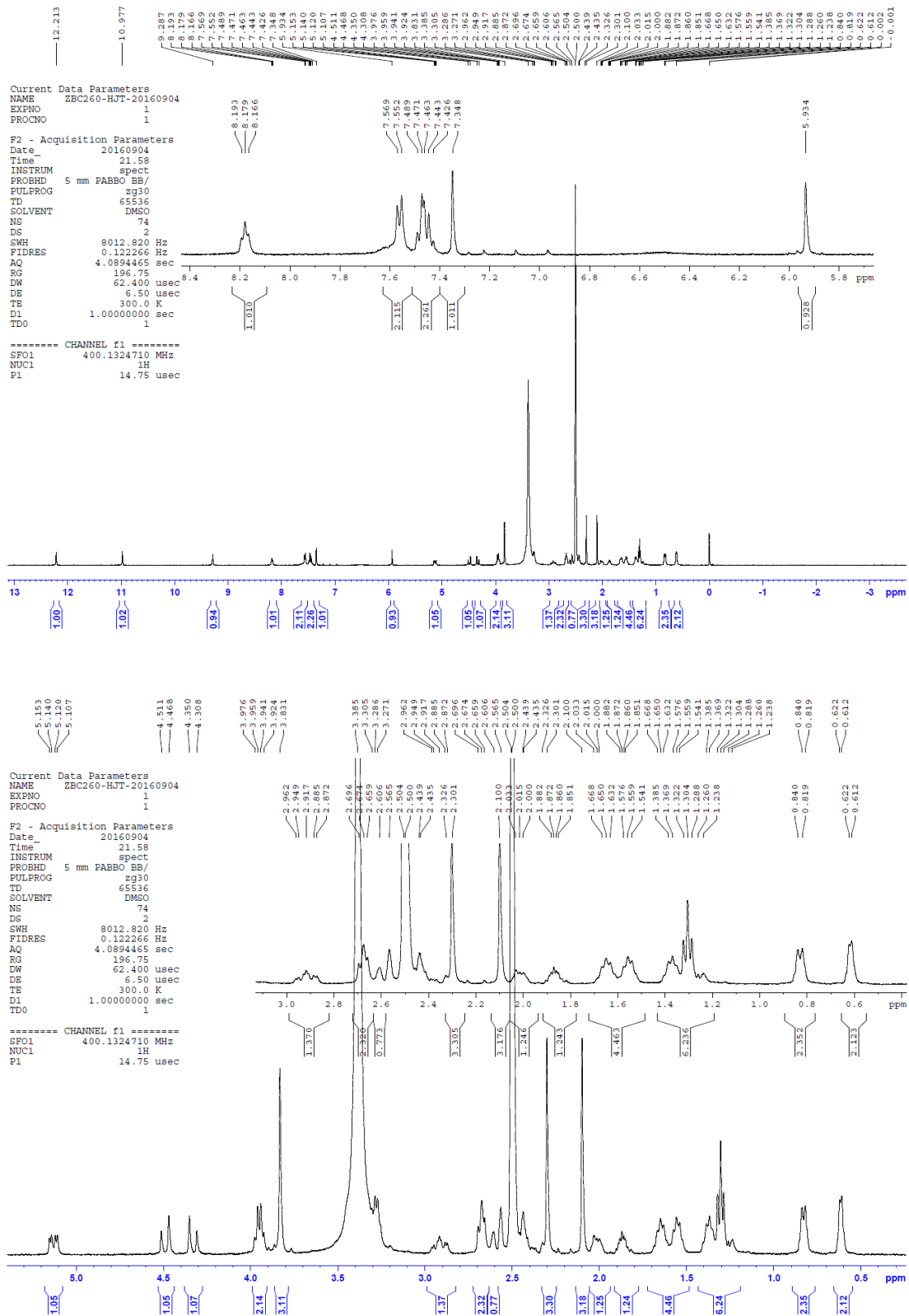
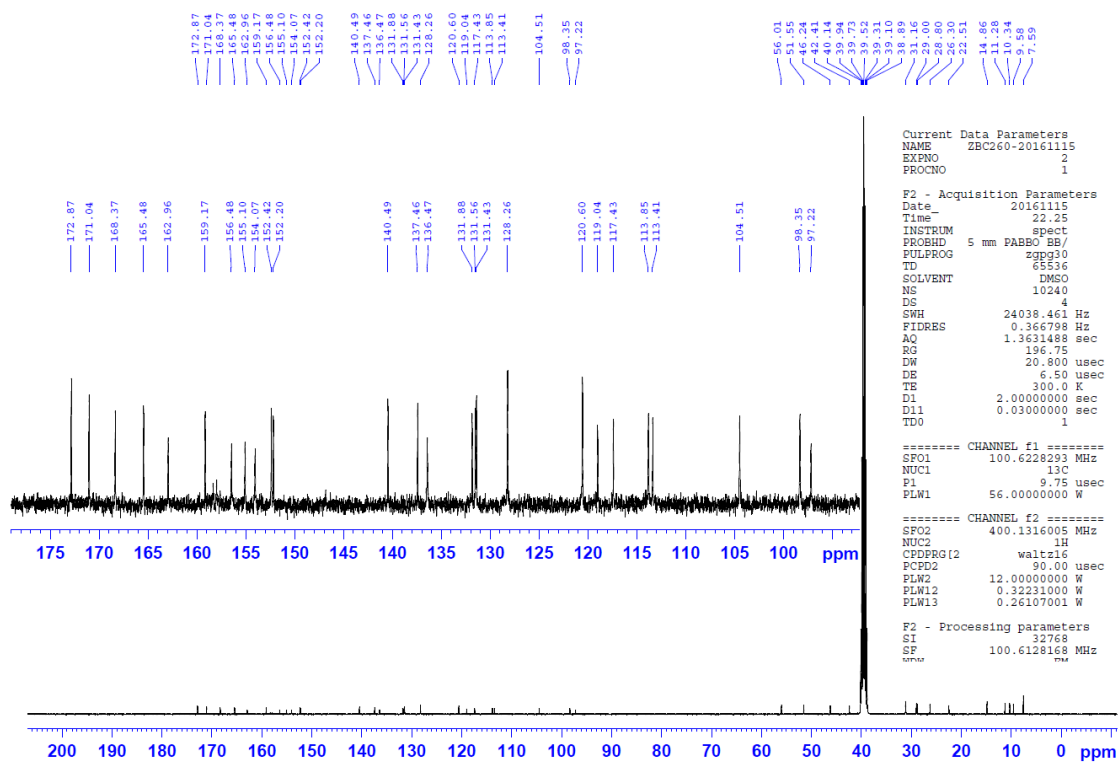
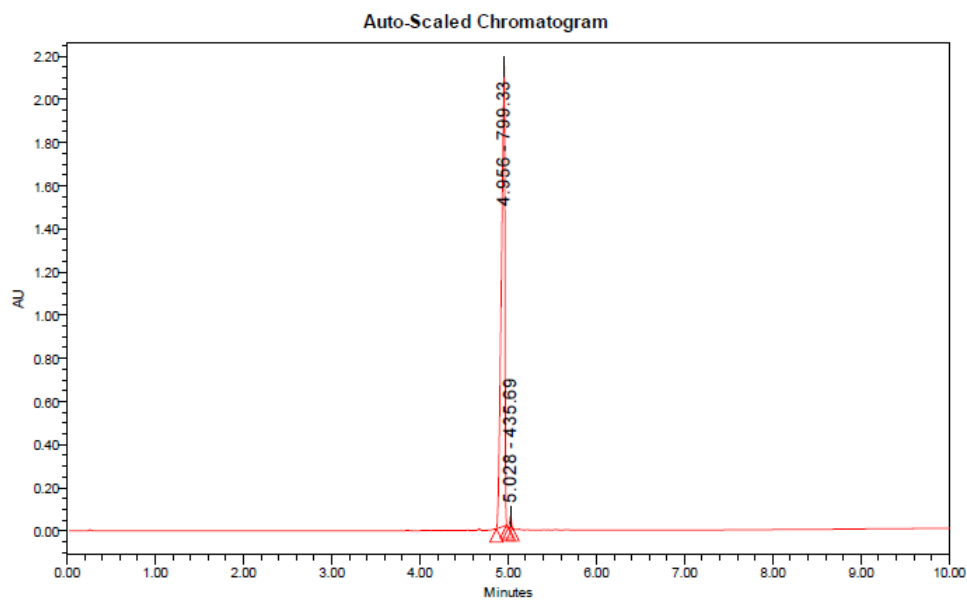


Figure S3.  $^{13}\text{C}$  NMR spectrum of compound **23**.



**Figure S4.** UPLC-MS results for compound **23**.

SAMPLE INFORMATION			
Sample Name:	HJT-A251-HPLC	Acquired By:	System
Sample Type:	Unknown	Date Acquired:	3/16/2016 6:19:33 PM EDT
Vial:	1:B,6	Acq. Method Set:	10to100% Bin 10
Injection #:	1	Date Processed:	3/16/2016 10:18 PM EDT
Injection Volume:	3.00 ul	Processing Method:	Caffeine PDA
Run Time:	10.0 Minutes	Channel Name:	254.0nm@2
Sample Set Name:	2	Proc. Chnl. Descr.:	PDA Spectrum (210-500)nm



**Peak Results**

	RT	Area	Height	% Area
1	4.956	5872907	2133708	98.85
2	5.028	68087	53990	1.15

**Peak Results**

	Base Peak (m/z)
1	799.33
2	435.69

**Figure S5.** Proposed metabolites of compound **23** in mouse liver microsomes

