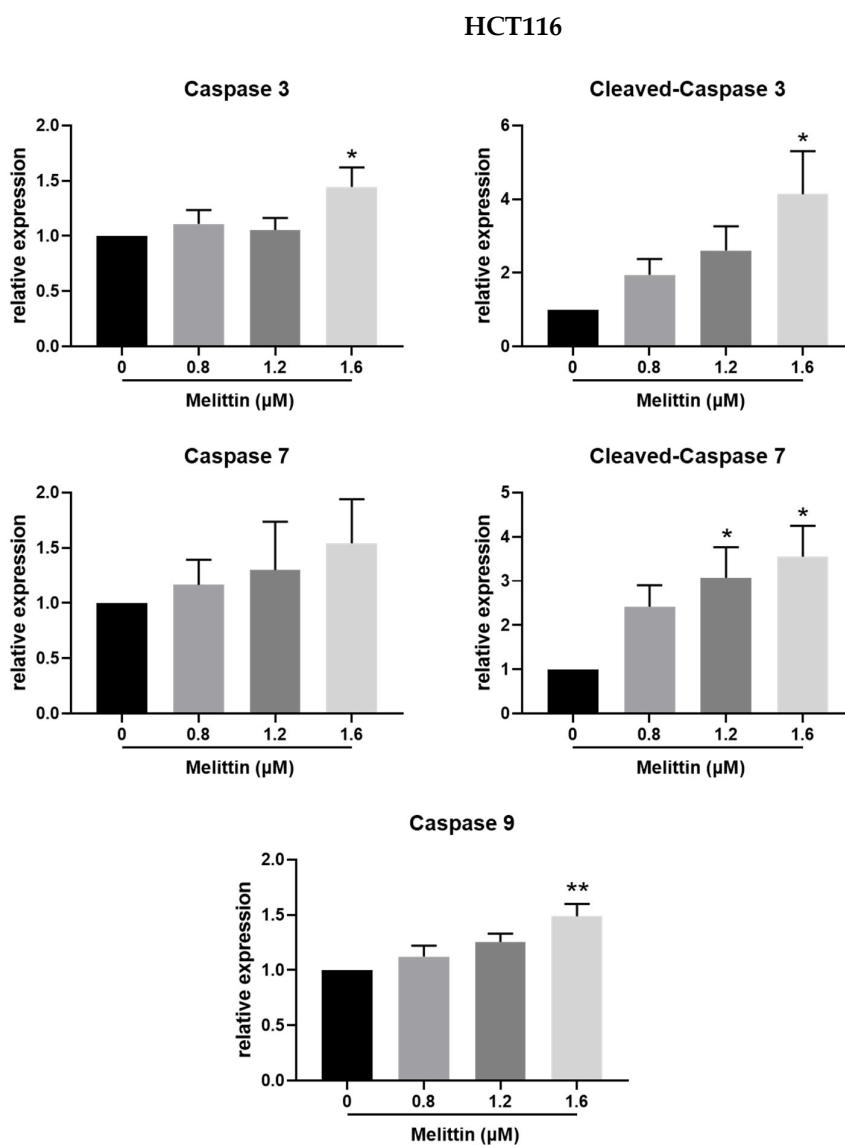


## Supplementary Information

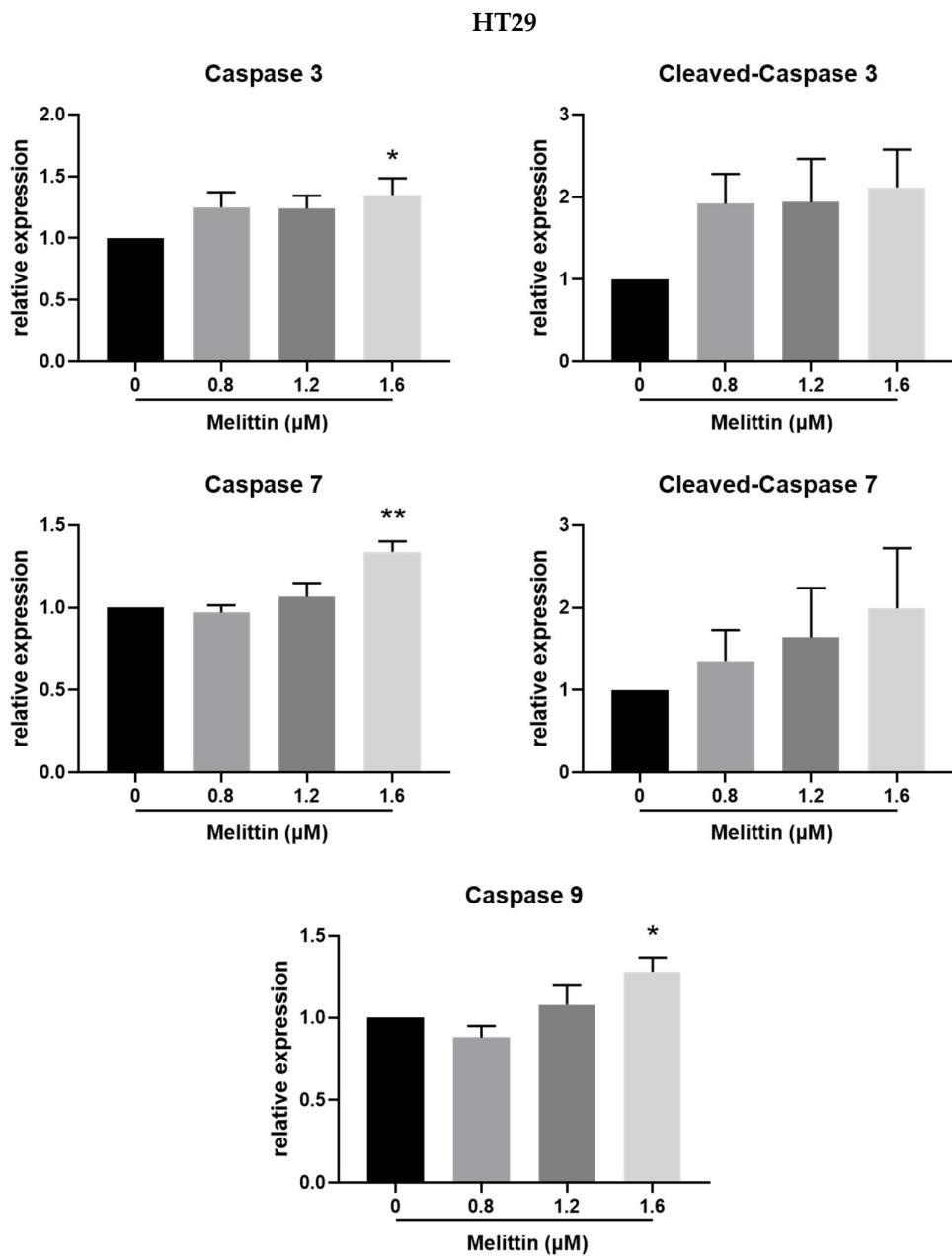
**Figure S1.** Melittin induces apoptosis in CRC cells by activating the caspase cascade. (A). The expression levels of Caspase 3, Cleaved-caspase 3, Caspase 7, Cleaved-caspase 7, and Caspase 9 in HCT116 cells; (B). The expression levels of Caspase 3, Cleaved-caspase 3, Caspase 7, Cleaved-caspase 7, and Caspase 9 in HT29 cells. Statistical significance: \*, p < 0.05, \*\*, p < 0.01, \*\*\*, p < 0.001.

A

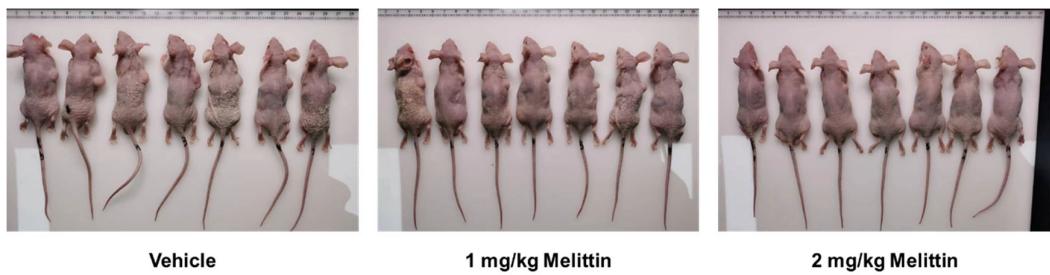


**Figure S1. continued**

B



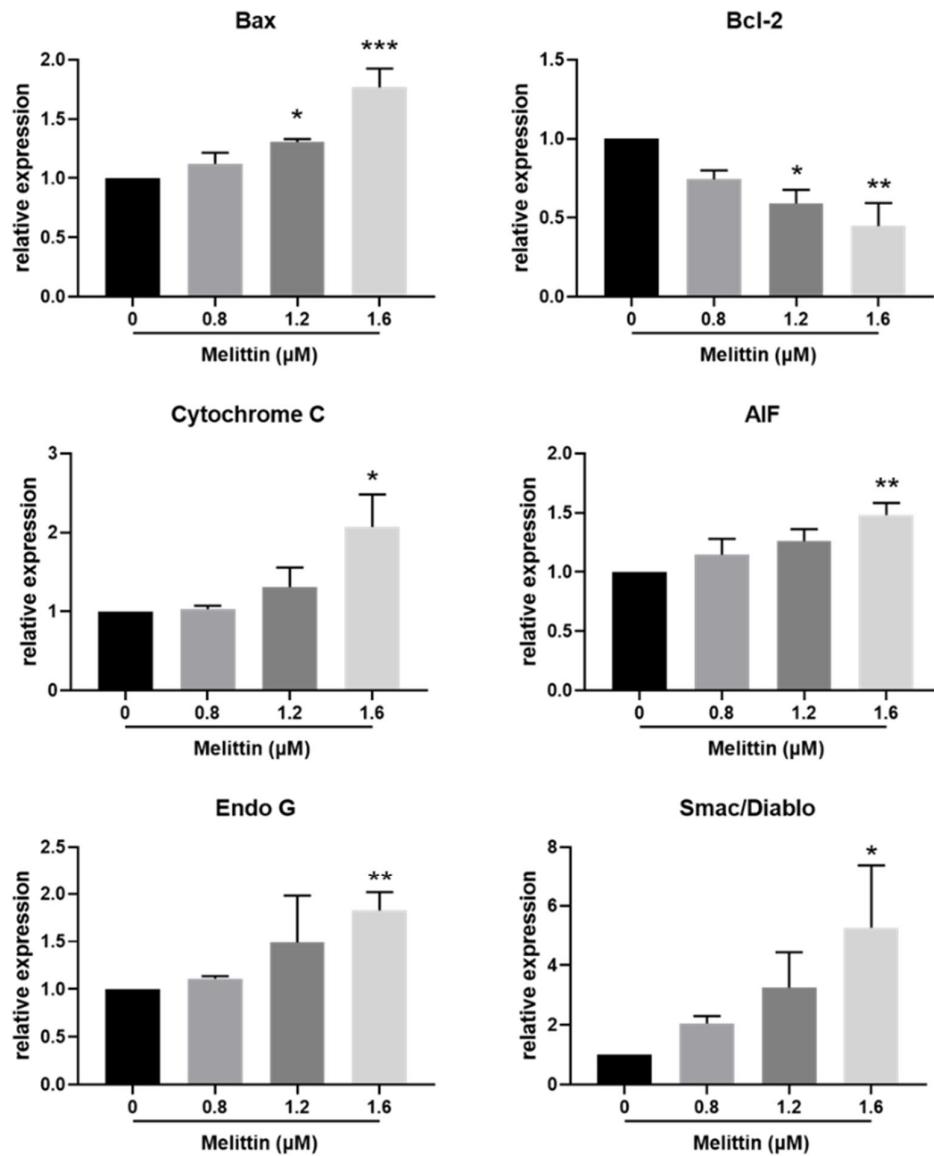
**Figure S2.** Melittin suppresses tumor growth in a subcutaneous heterograft tumor model using BALB/c nude mice. After treatment with melittin, the tumor volume in mice significantly decreased, with a more pronounced effect in the high-dose (2 mg/kg Melittin) group.



**Figure S3.** Melittin induces mitochondrial pathway-mediated apoptosis *in vitro*. (A). The expression levels of Bax, Bcl-2, Cytochrome C, AIF, Endo G, Smac/Diablo in HCT116 cells; (B). The expression levels of Bax, Bcl-2, Cytochrome C, AIF, Endo G, Smac/Diablo in HT29 cells. Statistical significance: \*, p < 0.05, \*\*, p < 0.01, \*\*\*, p < 0.001.

A

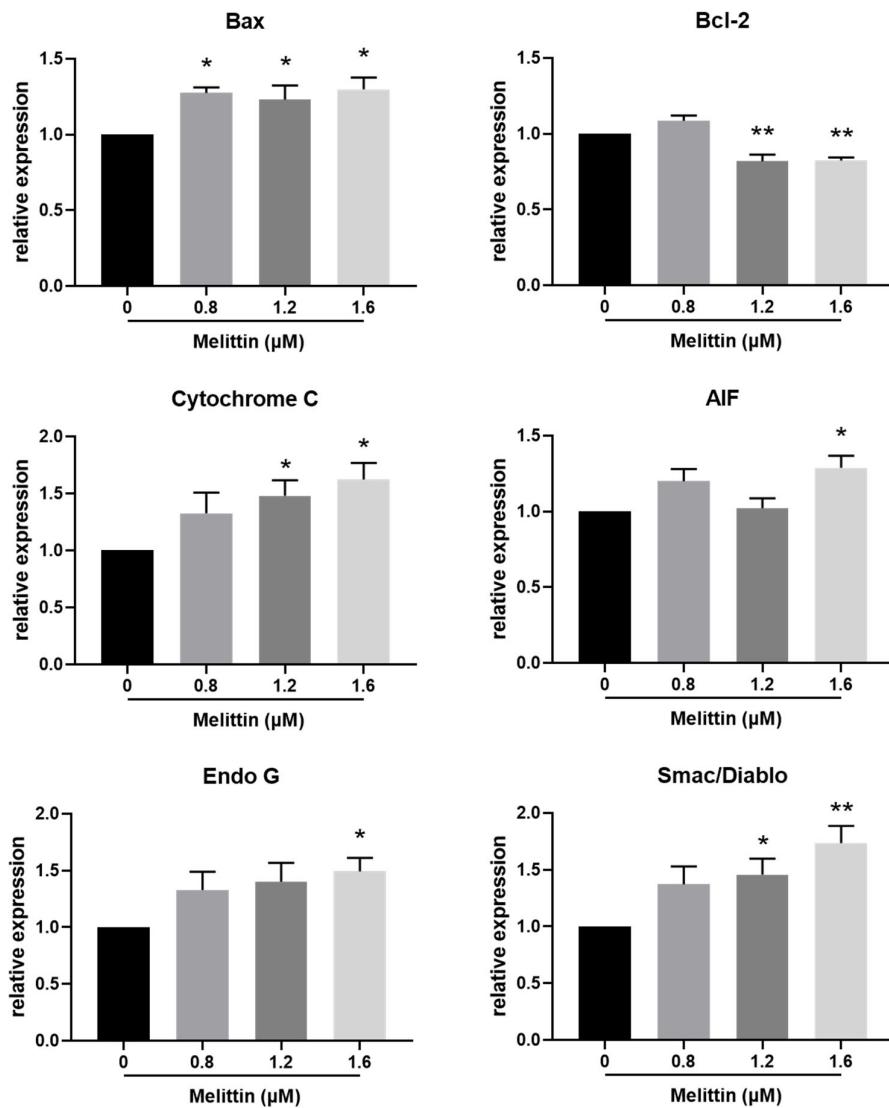
HCT116



**Figure S3. continued**

**B**

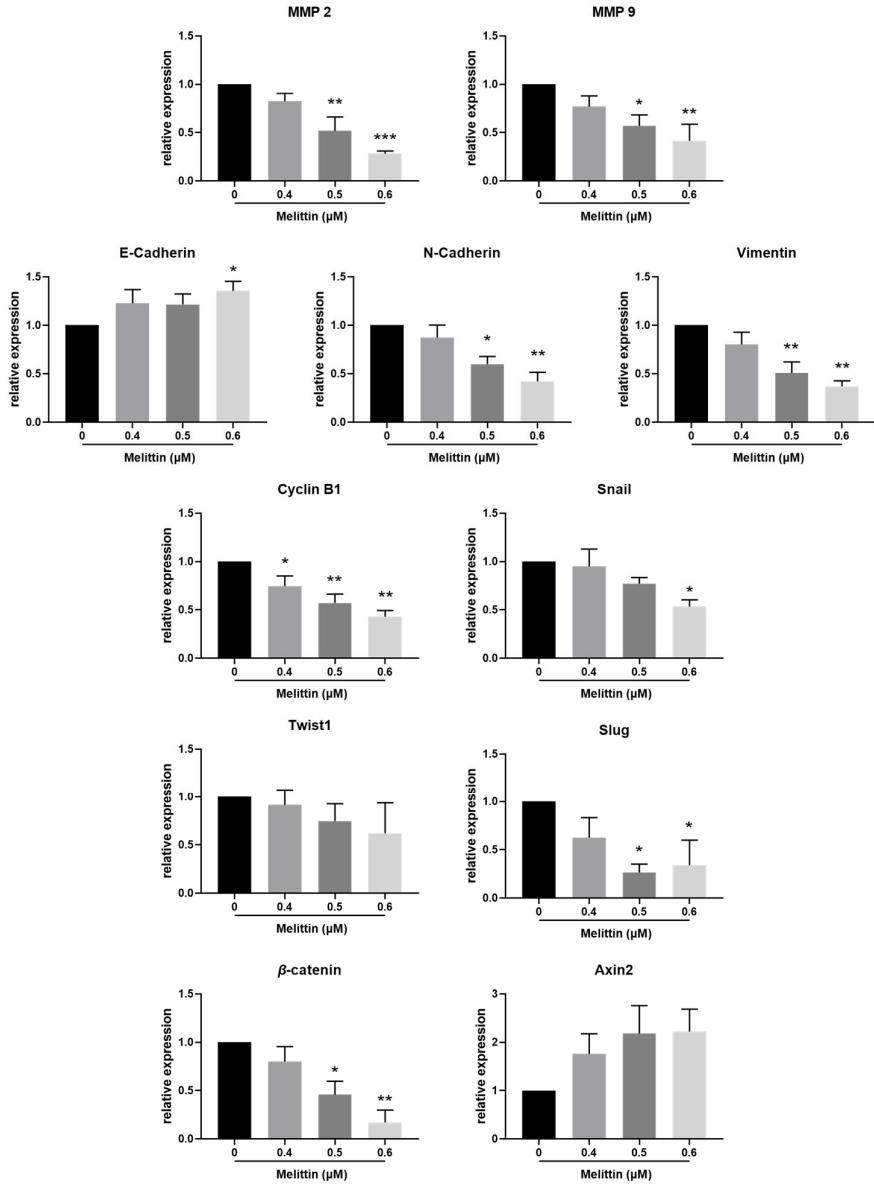
HT29



**Figure S4.** Melittin reverses EMT by inhibiting the  $\beta$ -catenin signaling pathway *in vitro*. (A). The expression levels of MMPs and EMT-related proteins in HCT116 cells; (B). The expression levels of MMPs and EMT-related proteins in HT29 cells. Statistical significance: \*, p < 0.05, \*\*, p < 0.01, \*\*\*, p < 0.001.

A

HCT116



**Figure S4. continued**

**B**

**HT29**

