

Supplementary Data

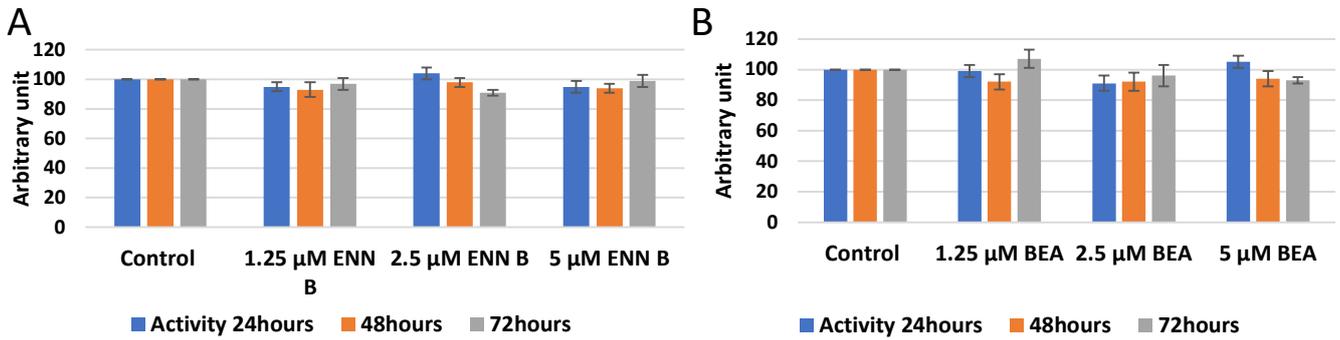


Figure S1. Impact of ENN B and BEA on proteolytic activities of cathepsin L in cellular extracts. **(A)** Cathepsin L activities were measured using cathepsin L specific substrate Z-F-R.AMC in cellular extracts after treatment with different ENN B and different periods concentrations. **(B)** Cathepsin L activities measured after treatments with different concentration for different periods with BEA. The data were analyzed using two-way ANOVA with Bonferroni's or Dunnett's post hoc analysis ($*p < 0.05$; $**p < 0.01$; $***p < 0.001$; $N = 7$). Statistics was calculated using GraphPad Prism.

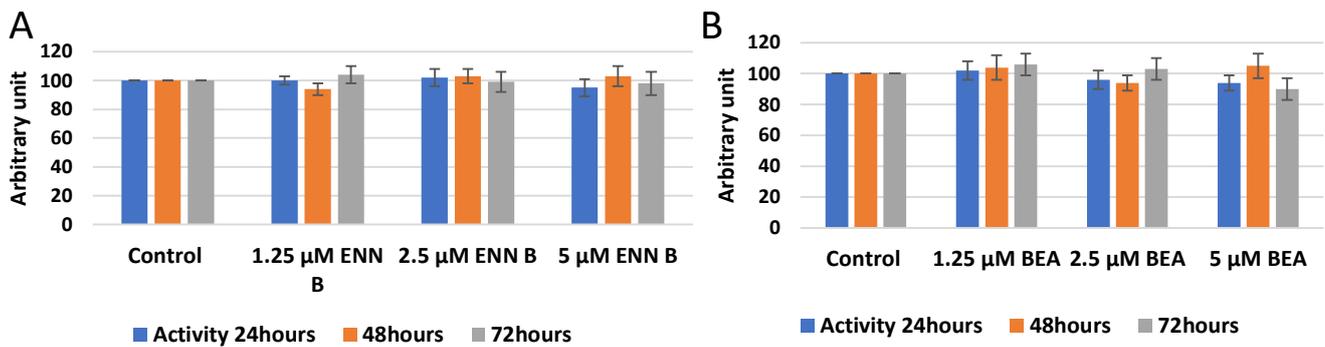
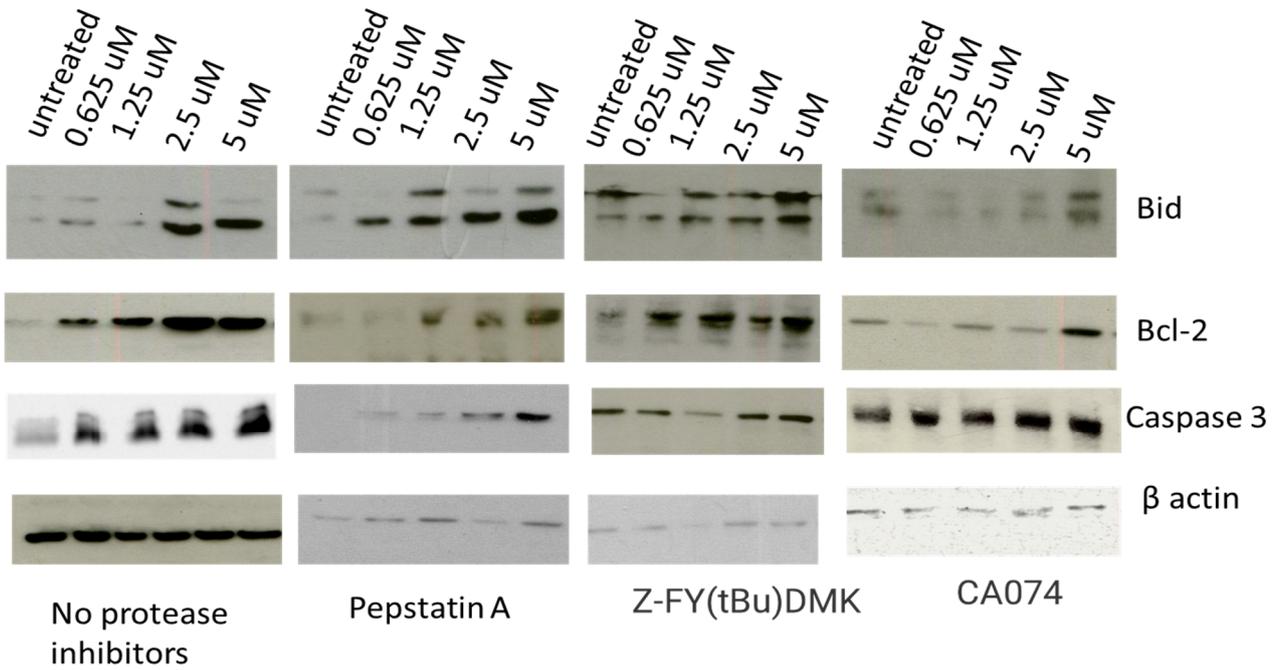
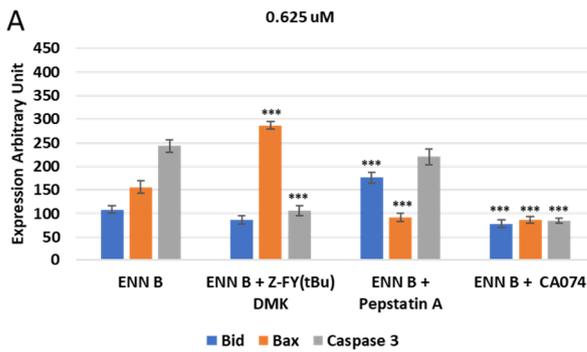


Figure S2. Impact of ENN B and BEA on proteolytic activities of Cathepsin L in media. **(A)** cathepsin L activities were measured using cathepsin L specific substrate Z-F-R.AMC in media after treatment with different ENN B and different concentrations for different periods. **(B)** cathepsin L activities measured in media after treatment with different concentrations of BEA for different periods. The data were analyzed using two-way ANOVA, Statistics was calculated using GraphPad Prism.

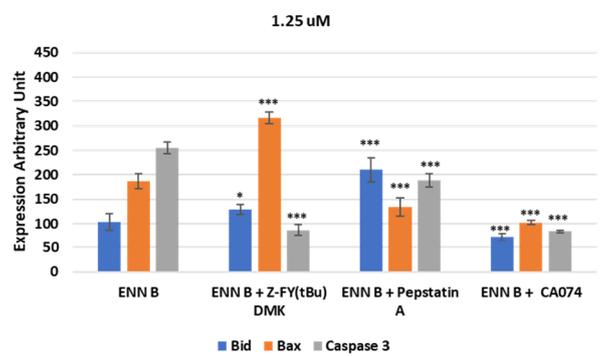
(1) ENN B



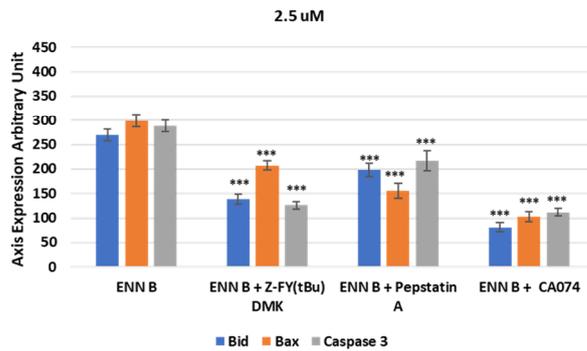
(2) A



C



B



D

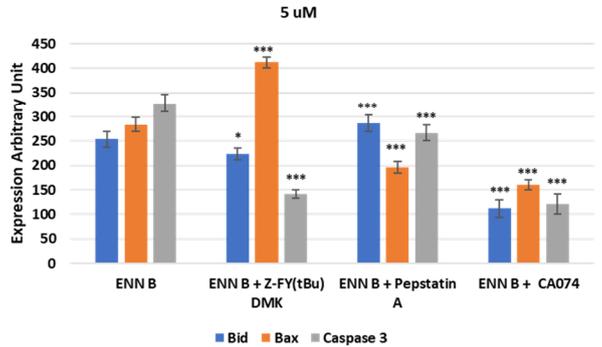


Figure S3. Impact of ENN B treatment on the expressions of some apoptotic proteins. (3.1) Western blot analysis of cleaved Bid, cleaved Bcl-2 and cleaved caspase 3 expression in KB-31- cells treated with 0, 0.625, 1.25, 2.5 and 5 μ M ENN B for 24 h in the presence and absence of 10 μ M of cathepsin D inhibitor pepstatin A or the same concentration of cathepsin L inhibitor Z-FY (tBu)DMK or cathepsin B inhibitor CA074. β -actin was used as an internal control. (3.2, A, B, C and D) Evaluation of cleaved Bid, cleaved Bcl-2 and cleaved caspase 3 expression after treatment with 0, 0.625, 1.25, 2.5 and 5 μ M ENN B for 24 h in the presence and absence of 10 μ M of cathepsin D, cathepsin L or cathepsin B specific inhibitors. The data were analyzed using two-way ANOVA with Dunnett's post hoc analysis (* p < 0.05; ** p < 0.01; *** p < 0.001; N = 7). Statistics was calculated using GraphPad Prism.

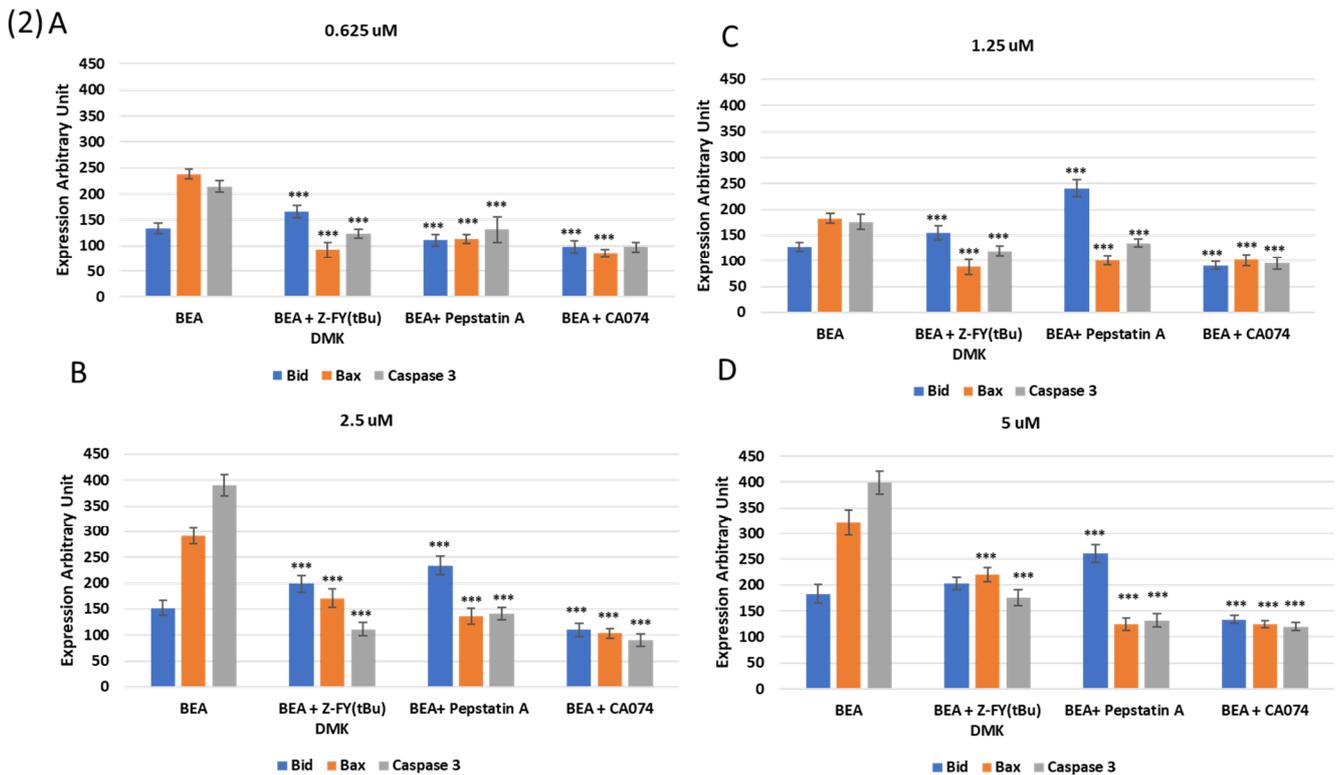
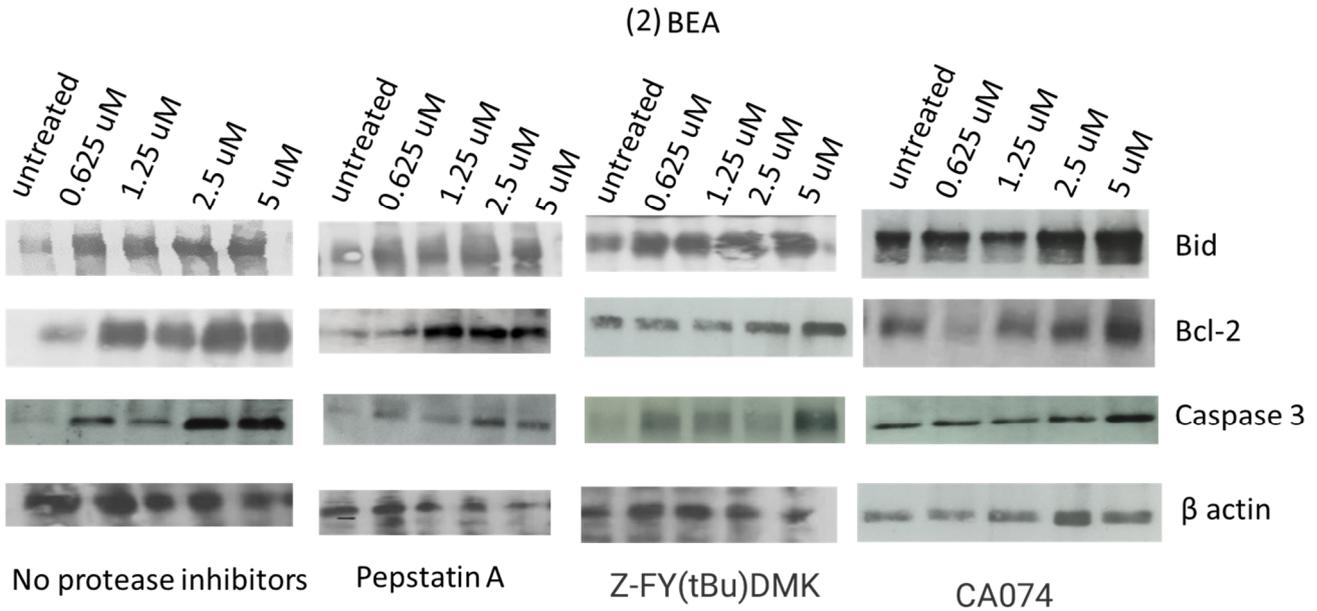


Figure S4. Impact of BEA treatment on the expressions of some apoptotic proteins. (4.1) Western blot analysis of cleaved Bid, cleaved Bcl-2 and cleaved caspase 3 expression in KB-31- cells treated with 0, 0.625, 1.25, 2.5 and 5 μ M BEA for 24 h in the presence and absence of 10 μ M of cathepsin D inhibitor pepstatin A or the same concentration of cathepsin L inhibitor Z-FY (tBu)DMK or cathepsin B inhibitor CA074. β -actin was used as an internal control. (4.2, A, B, C and D). Evaluation of cleaved Bid, cleaved Bcl-2 and cleaved caspase 3 expression after treatment with 0, 0.625, 1.25, 2.5 and 5 μ M BEA for 24 h in the presence and absence of 10 μ M of cathepsin D, cathepsin L or cathepsin B specific inhibitors. The data were analyzed using two-way ANOVA with Dunnett's post hoc analysis ($*p < .05$; $**p < .01$; $***p < .001$; $N = 7$). Statistics was calculated using GraphPad Prism.