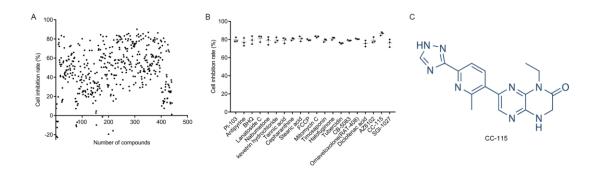
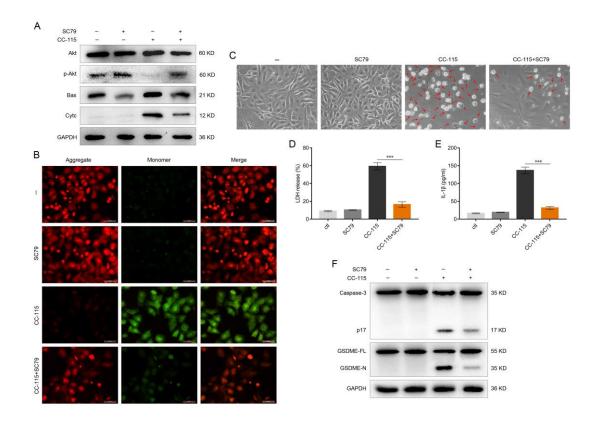
## **Supplementary materials**



**Figure S1.** Screening out CC-115 based on a library of 441 pyroptosis compounds. (**A**) A549 cells were treated with  $10 \,\mu\text{M}$  of the drug candidate library for 48 h. Each point represents the percentage of cell inhibition rate of the compounds. The cell viability was detected by CCK-8. (**B**) The top 20 compounds were confirmed in the secondary screening through CCK-8 assay. (**C**) Chemical structure of CC-115.



**Figure S2.** CC-115 promotes pyroptosis through inhibiting Akt/Bax signaling in H1650 cells. (**A**) Western blotting of Akt, Bax, and Cyt-c in H1650 cells treated with or without CC-115 (5 μM) and/or SC79 (5 μM). (**B**) Representative images of the mitochondrial membrane potential signal in H1650 cells determined using the JC-10 assay (scale bar, 50 μm). (**C**) Representative images of H1650 cells in the indicated groups (scale bar, 50 μm). (**D**, **E**) Release of LDH and IL-1β detected by ELISA. (**F**) Analysis of caspase-3 and GSDME expression using western blotting in the indicated groups. \*\*\*P < 0.001, two-tailed Student's t-test.