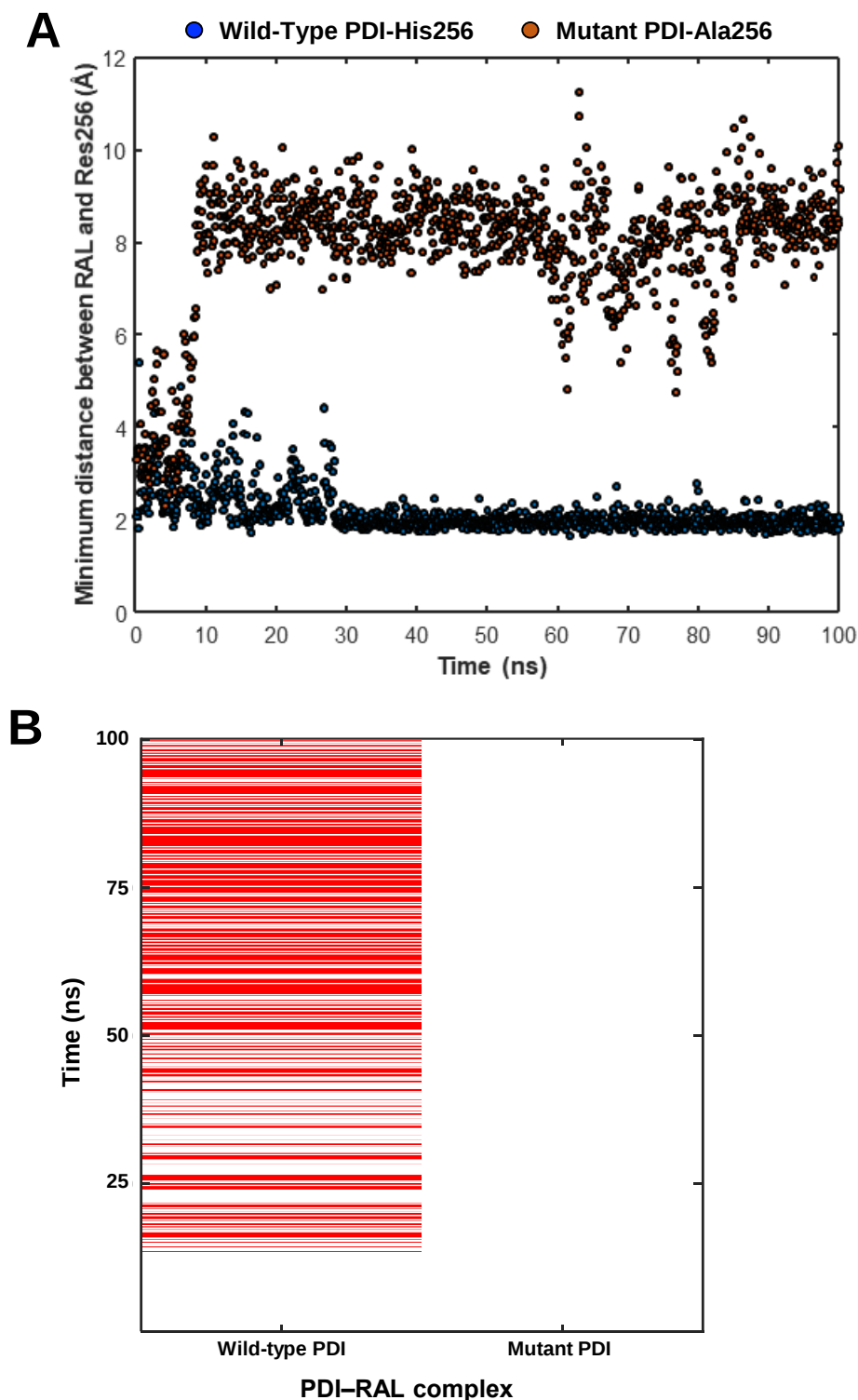
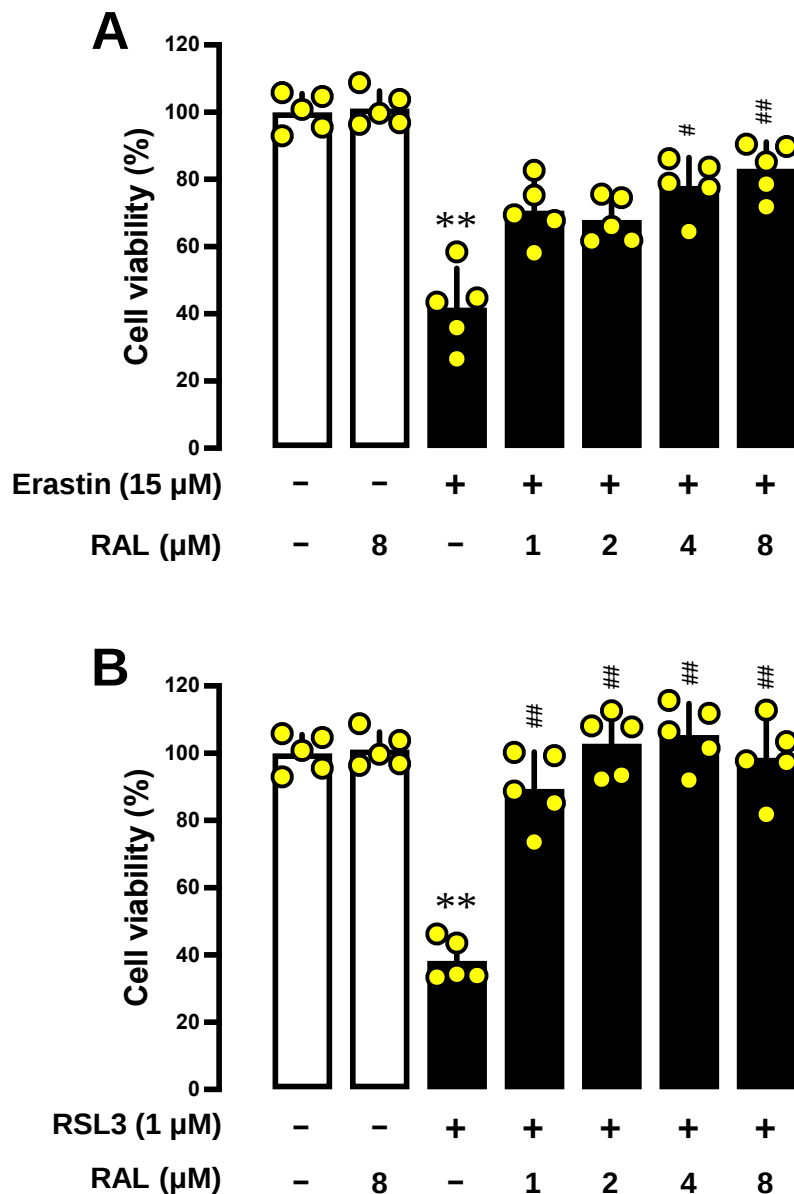


Supplemental Figure S1



Supplementary Fig. S1. Minimum distance (A) and the existence of a hydrogen bond (B) between PDI's residue-256 and RAL during the 100-ns MD simulation process. Note that the thickness of the red line in panel B reflects the time duration for the existence of the hydrogen bond between PDI and RAL. In the case of the wild-type PDI-His256-RAL complex, the hydrogen bond is present between PDI and RAL for a total of 60.1 ns during the whole 100-ns MD simulation process, whereas in the case of the mutant PDI-Ala256, the hydrogen bond is not present between PDI-Ala256 and RAL during the whole MD simulation process.

Supplementary Figure S2



Supplementary Fig. S2. RAL prevents erastin- or RSL3-induced ferroptosis in ER-negative MDA-MB-231 human breast cancer cells. **A.** Cells were treated with erastin $\pm$ RAL at the indicated concentrations for 24 h, and cell viability was determined by MTT assay ($n = 5$). **B.** Cells were treated with RSL3 $\pm$ RAL at the indicated concentrations for 24 h, and cell viability was determined by MTT assay ($n = 5$). All quantitative values are presented as mean $\pm$ S.D. (* or # $P < 0.05$; ** or ## $P < 0.01$).