Supplementary Materials: *Naja atra* Cardiotoxin 3 Elicits Autophagy and Apoptosis in U937 Human Leukemia Cells through the Ca²⁺/PP2A/AMPK Axis

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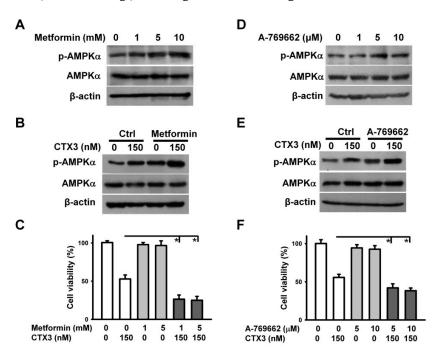


Figure S1. Effect of metformin and A-769662 on CTX3-induced cell death and AMPK α phosphorylation. Without specific indication, U937 cells were treated with 150 nM CTX3 for 4 h. (**A**) Effect of metformin on AMPK α phosphorylation in U937 cells. U937 cells were incubated with indicated metformin concentrations for 4 h; (**B**) Metformin enhanced CTX3-induced AMPK α phosphorylation. U937 cells were co-treated with 1 mM metformin and 150 nM CTX3 for 4 h; (**C**) Metformin enhanced CTX3-induced cell death. U937 cells were co-treated with indicated metformin and CTX3 concentrations for 4 h (mean ± SD, * p < 0.05); (**D**) Effect of A-769662 on AMPK α phosphorylation in U937 cells. U937 cells were incubated with indicated A-769662 concentrations for 4 h; (**E**) A-769662 enhanced CTX3-induced AMPK α phosphorylation. U937 cells were coincubated with 5 μM A-769662 and 150 nM CTX3 for 4 h; (**F**) A-769662 enhanced CTX3-induced cell death. U937 cells were incubated with indicated A-769662 and CTX3 concentrations for 4 h.

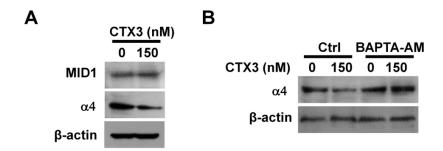


Figure S2. Effect of CTX3 on $\alpha 4$ and MID1 expression in CTX3-treated cells. Without specific indication, U937 cells were treated with 150 nM CTX3 for 4 h. On the other hand, U937 cells were pre-treated with 10 μ M BAPTA-AM for 1 h, and then incubated with 150 nM CTX3 for 4 h. (**A**) Western blot analyses of $\alpha 4$ and MID1 expression in CTX3-treated cells; (**B**) Effect of BAPTA-AM on $\alpha 4$ expression in CTX3-treated cells.