## **Supporting Information for**

## Batatasin III, a constituent of *Dendrobium scabrilingue*, improves murine pain-like behaviors with a favorable CNS safety profile

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## S1. Validation of LPS-induced inflammation in-vitro

LPS-induced inflammation models in RAW 264.7 macrophage and BV-2 microglial cells were validated using corresponding positive controls, dexamethasone, and minocycline, respectively. Cells were seeded in 24-well plates at  $2 \times 10^5$  cells/well density and incubated at 37°C for 24 h. Then the cells were treated with 50 µM of dexamethasone, minocycline, or batatasin III (BTS) at a concentration of 12.5, 25, and 50 µM for 1 h. LPS (1 µg/mL) was added to each well and incubated for 12 h in RAW 264.7 cells and 24h for BV-2 cells. Cell culture media was collected, and NO, IL-6, and TNF- $\alpha$  expression levels were analyzed. As shown in supplementary figure 1, dexamethasone inhibited the LPS-induced expression of NO, TNF- $\alpha$ , and IL-6 in RAW 264.7