Validation data for Isoliquiritigenin

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Isoliquiritigenin is an inhibitor of NF- κ B and the NLRP3 (NOD-like receptor (NLR) pyrin domain-containing protein 3) inflammasome. NF- κ B regulates multiple aspects of innate and adaptive immune functions and serves as a pivotal mediator of inflammatory responses. While the NLRP3 inflammasome is an innate immune sensor that is activated by a two-step process; a first signal ('priming') is provided by microbial molecules such as lipopolysaccharide (LPS), and the second signal is provided by a wide array of stimuli including endogenous molecules or crystalline substances such as monosodium urate (MSU) crystals. Inflammasome activation triggers caspase-1-mediated interleukin-1 β (IL-1 β) production and secretion.

The ability of isoliquiritigenin to inhibit the NLRP3 inflammasome was validated using InvivoGen's THP-1/HEK-Blue^M IL-1 β assay. This assay uses the secretion of IL-1 β by THP1-Null2 cells as an indicator of NLRP3 inflammasome induction. The IL-1 β production by these cells is measured using HEK-Blue^M IL-1 β cells. Treatment with isoliquiritigenin inhibited IL-1 β secretion in a dose-dependent manner (**Figure 1**).

Dose-dependent inhibition of NLRP3 activity

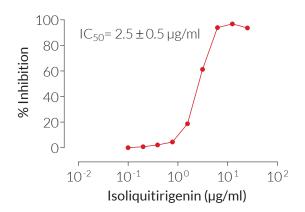


Figure 1: Isoliquiritigenin inhibits the NLRP3 inflammasome response in a dose-dependent manner.

THP1-Null2 cells, primed with LPS-EK (1 μ g/ml for 3 h), were stimulated with MSU (150 μ g/ml) and increasing concentrations of isoliquiritigenin. After overnight incubation, IL-1 β secretion was analyzed by adding 50 μ l of supernatant from treated THP1-Null2 cells to HEK-Blue^M IL-1 β cells. IL-1 β induced activation of NF- κ B was assessed by measuring the levels of SEAP in the supernatant of HEK-Blue^M IL-1 β cells using QUANTI-Blue^M Solution, a SEAP detection reagent, and by reading the optical density (OD) at 655 nm. Data are shown as a percentage (%) inhibition of the maximal response for the ligand with no inhibitor.



InvivoGen Hong Kong: +852 3622-3480 E-mail: info@invivogen.com