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SUPPLEMENTAL FIGURE LEGENDS

<u>Supplemental Fig. 1.</u> Expression of IL-1R2 in cancer cell lines. (A,B) Cancer cell lines from the panel were treated with 1ng/mL IL-1 β for 24h. Proteins were isolated and Western immunoblotted with antibodies recognizing IL-1R2, cIAP1, cIAP2 and XIAP. β -actin was used as a loading control.

<u>Supplemental Fig. 2.</u> Dosage response of cancer cell lines to SM-164 and IL-1 β treatment. (A) SF295, (C) SNB75 and (E) U2OS cells were treated with 1ng/mL IL-1 β in the presence of varying concentrations of SM-164 for 48h. (B) SF295, (D) SNB75 and (F) U2OS cells were treated with 100n SM-164 in the presence of varying concentrations IL-1 β of for 48h. Cell viability was determined by Alamar Blue. The percentage viability relative to vehicle +/- SD (n = 4) was plotted.

<u>Supplemental Fig. 3.</u> SMC treatment does not impair IL-1β-mediated activation of the NF- κ B pathway. (A) SNB75 and (B) U2OS cells were treated with vehicle (veh) or 100nM SM-164 (SMC) for 2h, then treated with 1ng/mL IL-1β for the indicated times, and protein extracts were immunoblotted for I κ B α , p65-pS536, p65, cIAP1, cIAP2 and XIAP. β-actin was used as a loading control. Asterisk denotes a non-specific band.

<u>Supplemental Fig. 4.</u> Expression of dominant negative IKK β blocks IL-1 β -mediated activation of the NF- κ B pathway. (A) SNB75 and (B) U2OS cells were transduced with adenoviral vectors expressing GFP or co-expressing GFP and a dominant negative form of IKK β (IKK β -DN) at an MOI of 100 (SNB75) or 50 (U2OS) for 24h. Cells were

treated with vehicle (veh) or 100nM SM-164 (SMC) for 2h, and then treated with 1ng/mL IL-1 β for 10min. Cells were harvested and the protein expression levels of I κ B α , IKK β , cIAP1, cIAP2 and XIAP were analysed by Western immunoblotting. β -actin was used as a loading control. Asterisk denotes a non-specific band.

<u>Supplemental Fig. 5.</u> LPS synergizes with SM-164 to induce cancer cell death. SF295, SNB75 and U2OS cells were treated with combinations of $1\mu g/mL$ LPS and 100nM SM-164 for 48h. Cell viability was measured with Alamar Blue. Percentage viability relative to vehicle +/- SD (n = 4) was plotted.