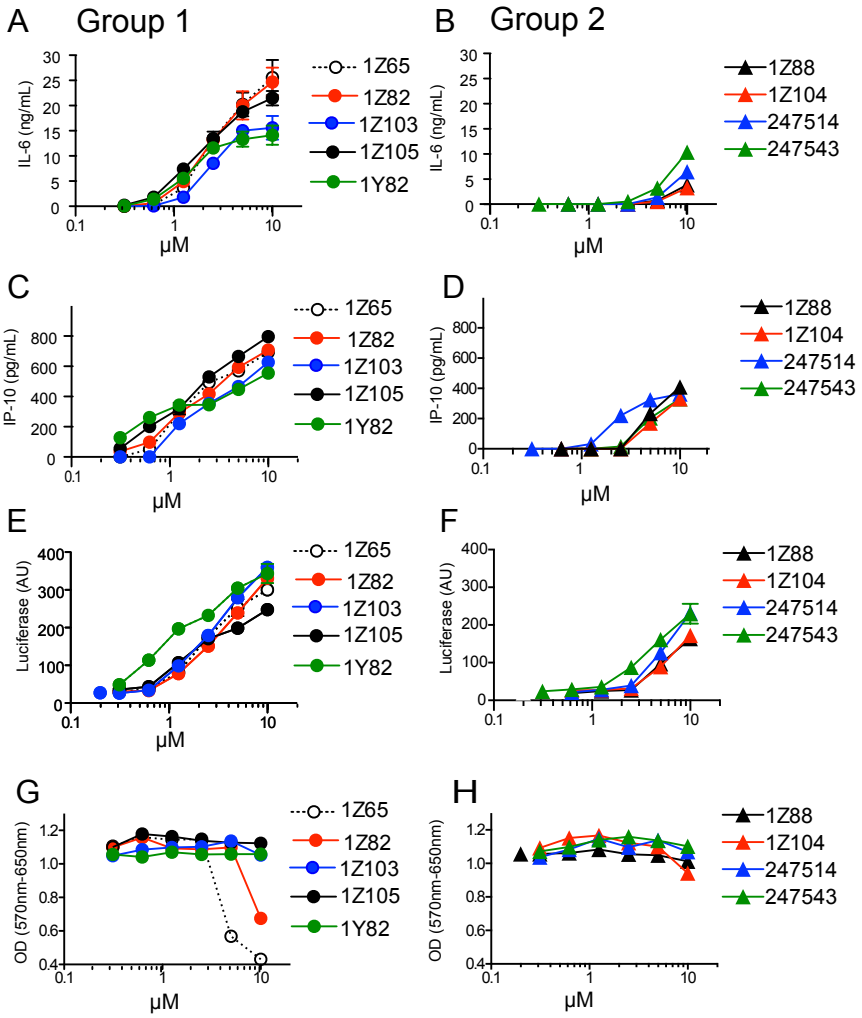
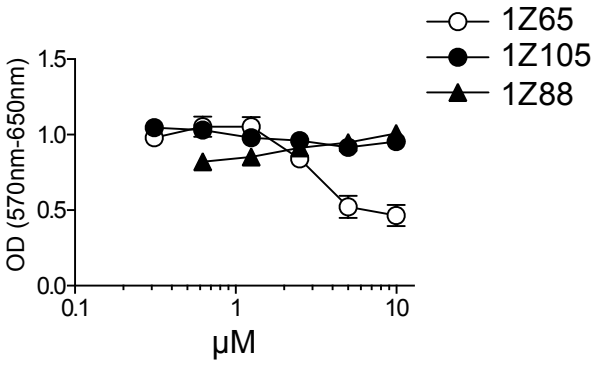
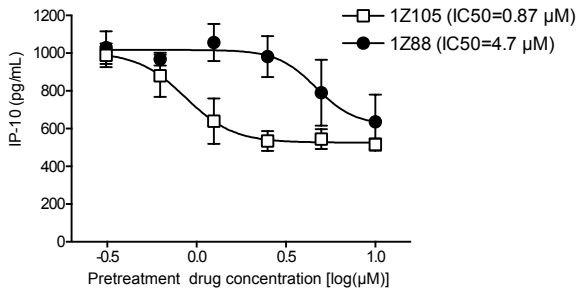


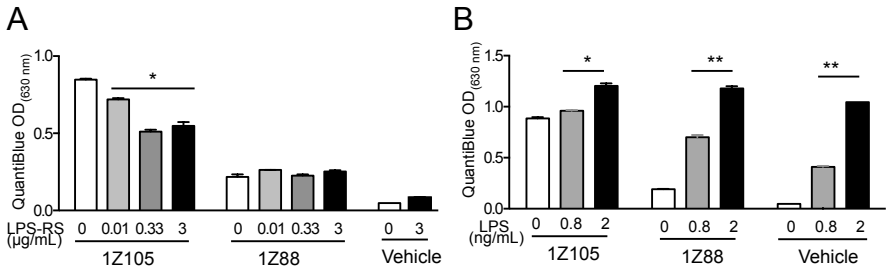
Supplemental Figure 1. Dose dependent cytokine production and toxicity. WT C57BL/6 BMDC were plated at 1 ×105 cells/well and were incubated with graded concentrations of each compound overnight. IL-6 (A and B) and IP-10 (C and D) levels were measured by ELISA. Type 1 IFN levels were measured by IRSE reporter assay (E and F). Control wells included LPS (10ng/mL) or MPLA (1 μg/mL). Induction of IL-6 and IP-10 by LPS was 17.3 ± 2.4 ng/mL and 387 ± 16 pg/mL , respectively. Induction of IL-6 and IP-10 by MPLA were 3.8 ± 0.2 ng/mL and 106 ± 39 pg/mL, respectively. The type 1 IFN level induced by MPLA (1 μg/mL) was 53 ± 1.5 AU. Hep G2 cells in 96-well plates were exposed to compounds for 24 hours and cell viabilities were measured by a tetrazolium (MTT) assay (G and H). The OD(570-650) was 1.14 ± 0.01 for the vehicle control treated wells. All data shown are mean ± SEM of triplicates and representative of two independent experiments.



Supplemental Figure 2. Dose dependent toxicity of 1Z65, 1Z105 and 1Z88. WT C57BL/6 BMDC were plated at 1 $\times 10^5$ cells/well and were incubated with graded concentrations of each compound overnight. Cell viabilities were measured by MTT assay. The OD(570-650) was 0.98 ± 0.03 for the vehicle control treated wells. All data shown are mean ± SEM of triplicates and representative of two independent experiments.



Supplemental Figure 3. Desensitization of IP-10 release by BMDM by pretreatment with 1Z105 and 1Z88. WT C57BL/6 BMBM were plated at 5 $\times 10^4$ cells/well and were incubated with graded concentrations of 1Z105 or 1Z88 overnight. The cells were then challenged with LPS (10 ng/mL) and cytokine release in the culture supernatants after 18 h was assessed by ELISA. Data shown are mean ± SEM.



Supplemental Figure 4. Costimulation of LPS-RS or LPS with 1Z105 or 1Z88.
Murine TLR4-NF-κB reporter cells were incubated with 1Z105(5μM) or 1Z88 (10 μM) in the presence and absence of indicated concentrations of LPS-RS (A) or LPS (B) overnight. NF-κB activation was measured by SEAP release (OD_{630}). Data shown are mean ± SEM.
* $p<0.05$; **$p<0.01$ by one way ANOVA compared to cells without LPS-RS oe LPS with Dunnet's post hoc testing.