

Figure S1. NKG2D deficiency alters NOD CTL cytokine production but not lytic granule release. (A) Fold change (mean \pm SEM) in TNF α , IFN γ and IL-10 secretion by *Klrk1*^{-/-} CTL compared with wild type CTL. **(B)** Fold change (mean \pm SEM) in percent of CD107a⁺ *Klrk1*^{-/-} CTL compared with percent of CD107a⁺ wild type CTL. Data are combined from at least 4 independent experiments. * $p \leq 0.05$, ** $p \leq 0.01$, *** $p \leq 0.001$, **** $p \leq 0.0001$ in two-tailed Wilcoxon test.

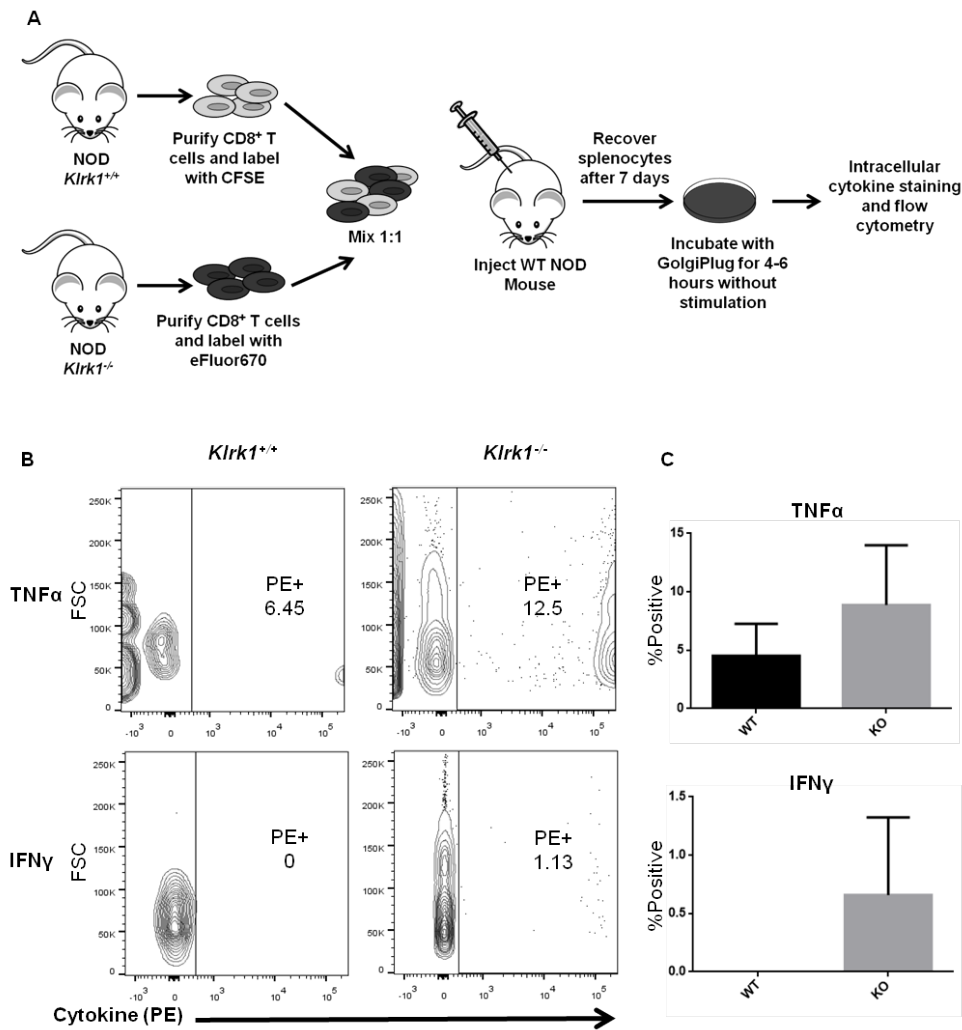


Figure S2. *Klrk1*^{-/-} NOD CTL produce increased TNF α and IFN γ in antibiotic-treated mice.

(A) Schematic of adoptive transfer and intracellular cytokine staining of labeled CD8⁺ T cells from wild type and *Klrk1*^{-/-} mice. **(B)** Representative flow cytometry plots showing intracellular staining for TNF α and IFN γ in *Klrk1*^{-/-} and wild type NOD CD8⁺ T cells one week after co-transfer into wild type ABX-treated NOD adoptive transfer recipient mice. **(C)** Combined results of two independent experiments (mean \pm SEM) showing percent TNF α or IFN γ positive *Klrk1*^{-/-} and wild type NOD CD8⁺ T cells 1 week after co-transfer into wild type ABX-treated NOD adoptive transfer recipient mice.

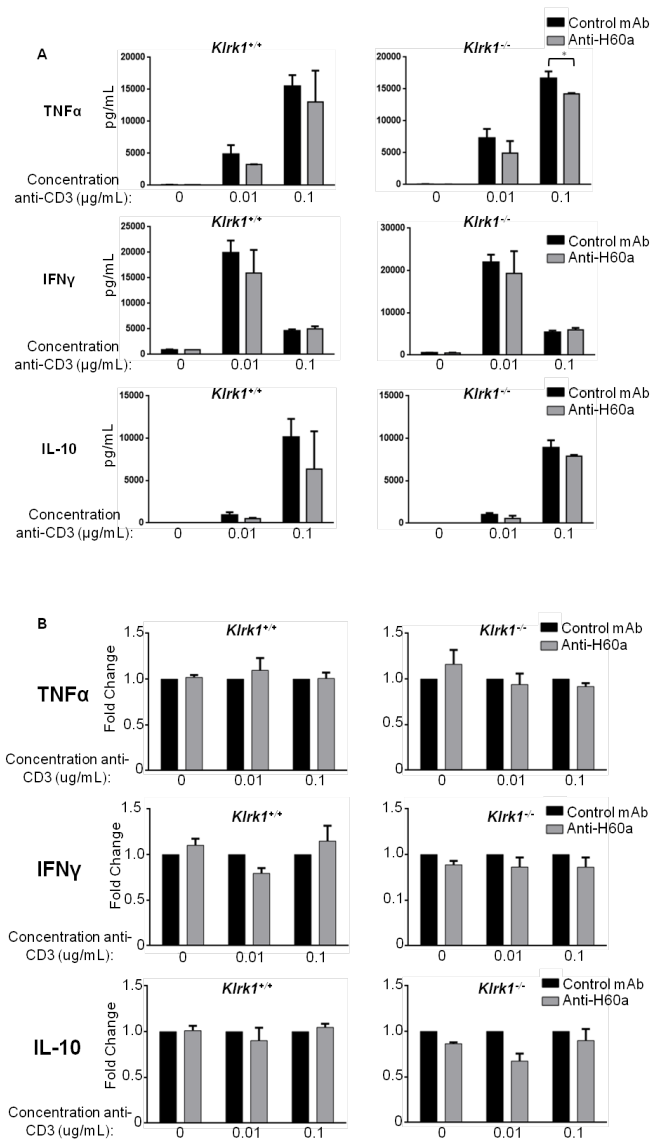


Figure S3. CTL effector cytokine production is not affected by blockade of NKG2D-H60a interaction during NOD CTL effector response. (A) Representative experiment showing TNF α , IFN γ and IL-10 secretion (mean \pm STD) by *Klrk1^{-/-}* and wild type CTL stimulated with anti-CD3 ϵ in the presence of an anti-H60a or isotype control antibody. Data are representative of at least 4 independent experiments. * $p \leq 0.05$ in one-tailed unpaired Mann-Whitney test. **(B)** Fold change (mean \pm SEM) in TNF α , IFN γ and IL-10 secretion by CTL stimulated with anti-CD3 ϵ in the presence of anti-H60a compared with isotype control antibody. Data are combined from at least 4 independent experiments.

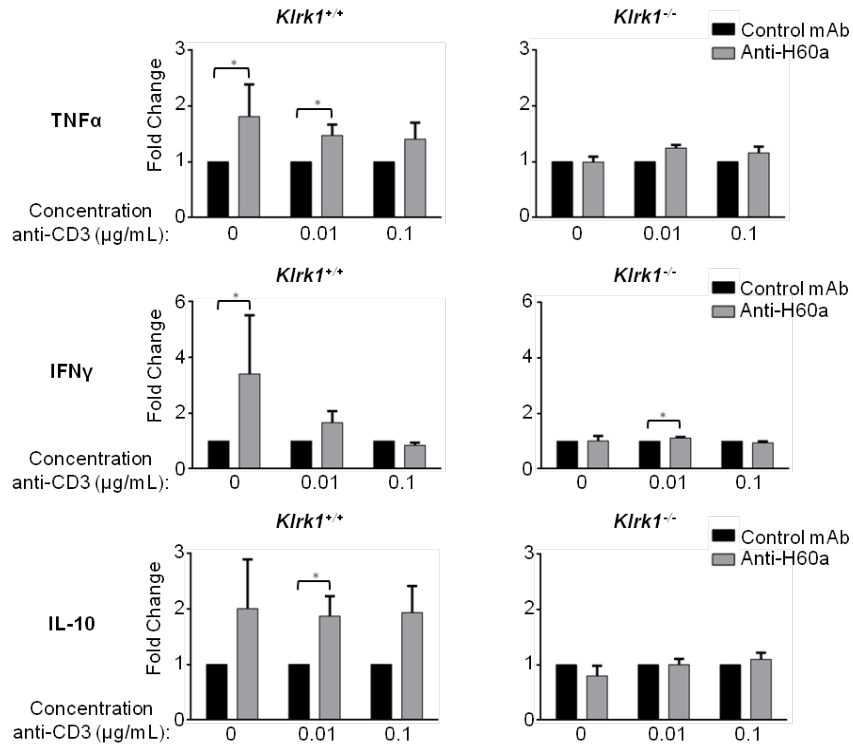


Figure S4. Blockade of NKG2D-H60a interaction during NOD CTL differentiation increases CTL effector cytokine production. Fold change (mean \pm SEM) in TNF α , IFN γ and IL-10 secretion by *Klrk1*^{-/-} and wild type CTL stimulated with anti-CD3 ϵ generated in the presence of an H60a blocking compared with isotype control antibody. Data are combined from at least 6 independent experiments. * $p \leq 0.05$ in one-tailed Wilcoxon test.