

B.

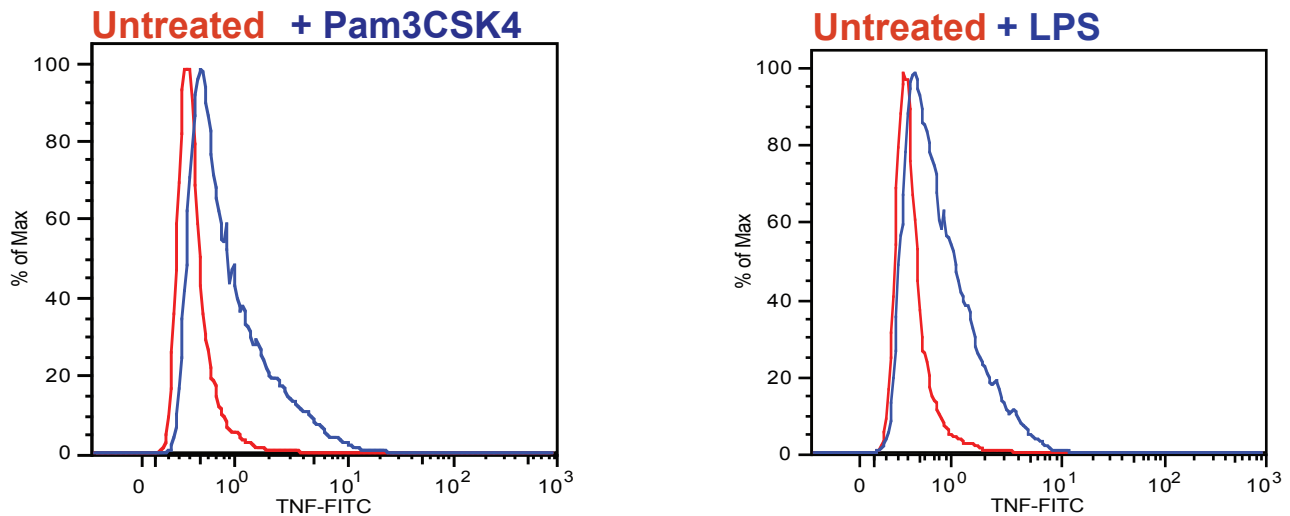
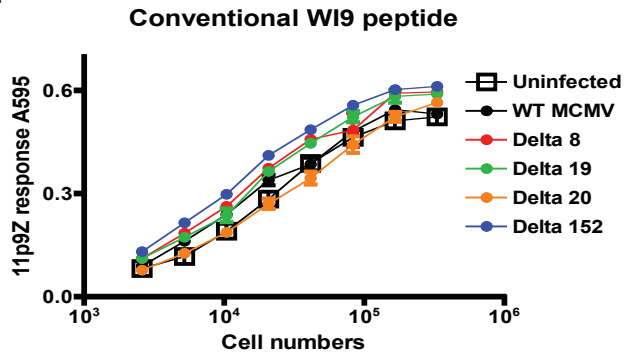


Fig S1 A: Relative T-cell responses to the cryptic peptide are significantly enhanced upon TLR-ligand stimulation, compared to the conventional peptide. T-cell responses to the WI9 and LYL8 peptides, upon Pam3CSK4, CpG, LPS, stimulation, were normalized to that of the untreated samples for 3 distinct experiments. An unpaired t-test (with Welch's correction) was performed, comparing the conventional and cryptic peptide responses. p-value < 0.05

B: Toll-like receptor agonists stimulated TNF- α production in macrophages: Primary macrophages from the WI9.LYL8 mice were stimulated with either LPS (1 μ g/mL) or Pam3CSK4 (1 μ g/mL) for a period of 6 hours, after which the macrophages were stained intracellularly for TNF- α and analyzed by flow cytometry.

A i.



ii.

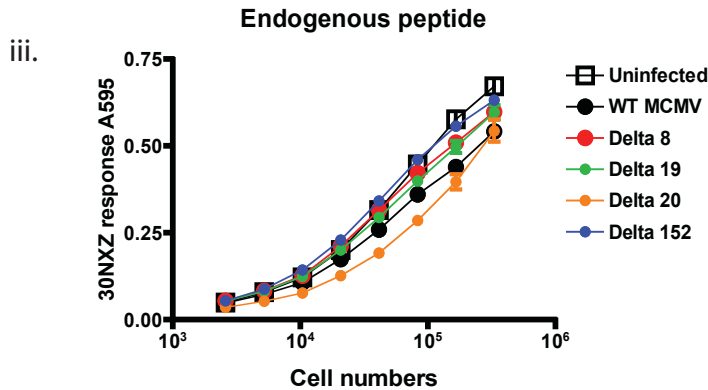
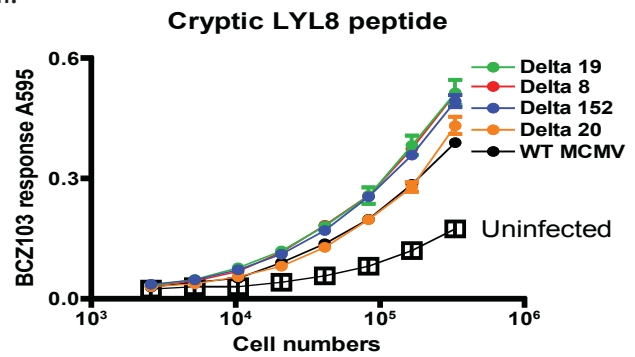


Fig S2A: Infection with various MCMV deletion mutants also enhances cryptic peptide presentation:

WI9.LYL8 macrophages were infected wild-type MCMV and MCMV mutants lacking multiple open-reading frames of the MCMV genome. The mutant Delta 152 indicates that it lacks the m152 gene that encodes for MHC Class I inhibitors. The cells were harvested and incubated with either the 11p9Z, BCZ103 or the 30NXZ hybridoma that is specific for an endogenous peptide. Data is representative of 2 independent experiments.

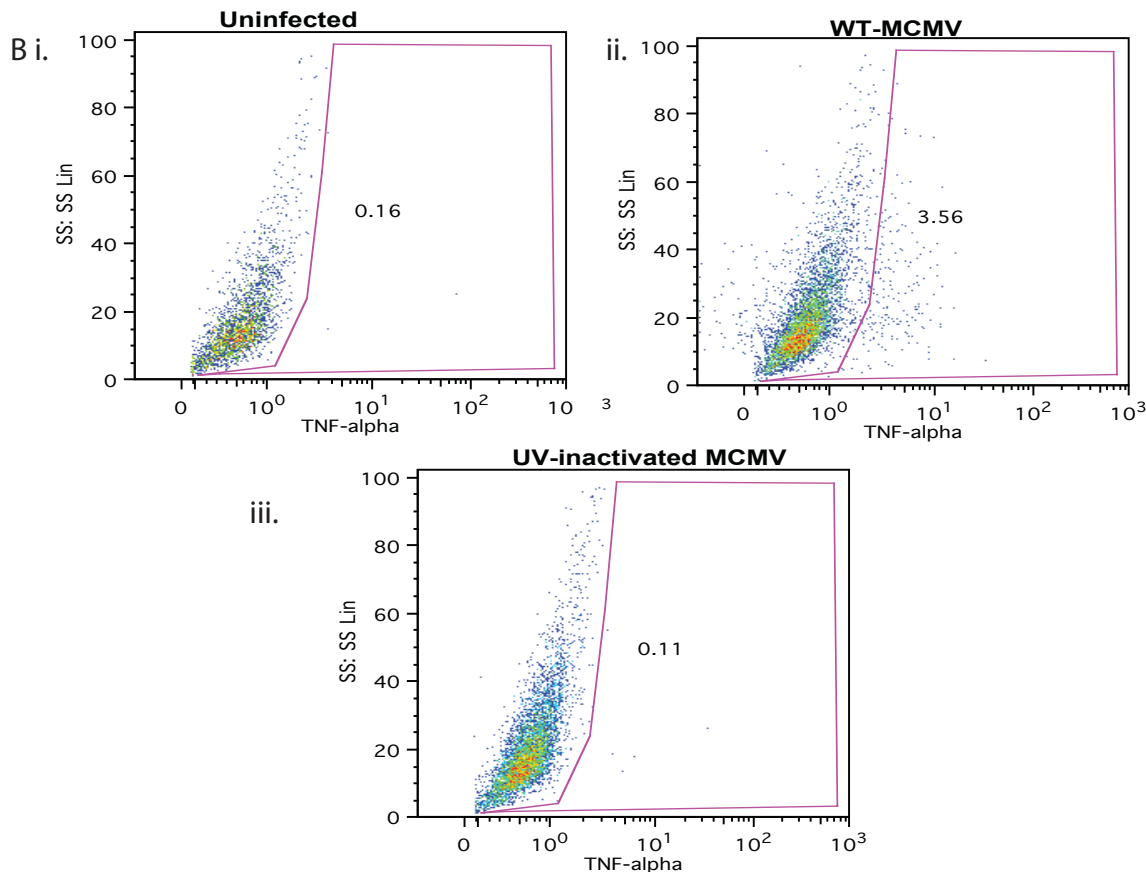


Fig S2B: UV-inactivated MCMV does not induce inflammatory cytokine production in macrophages: Uninfected and MCMV infected macrophages were stained intracellularly with TNF- α antibody and analyzed by flow cytometry. Data is representative of 3 experiments.

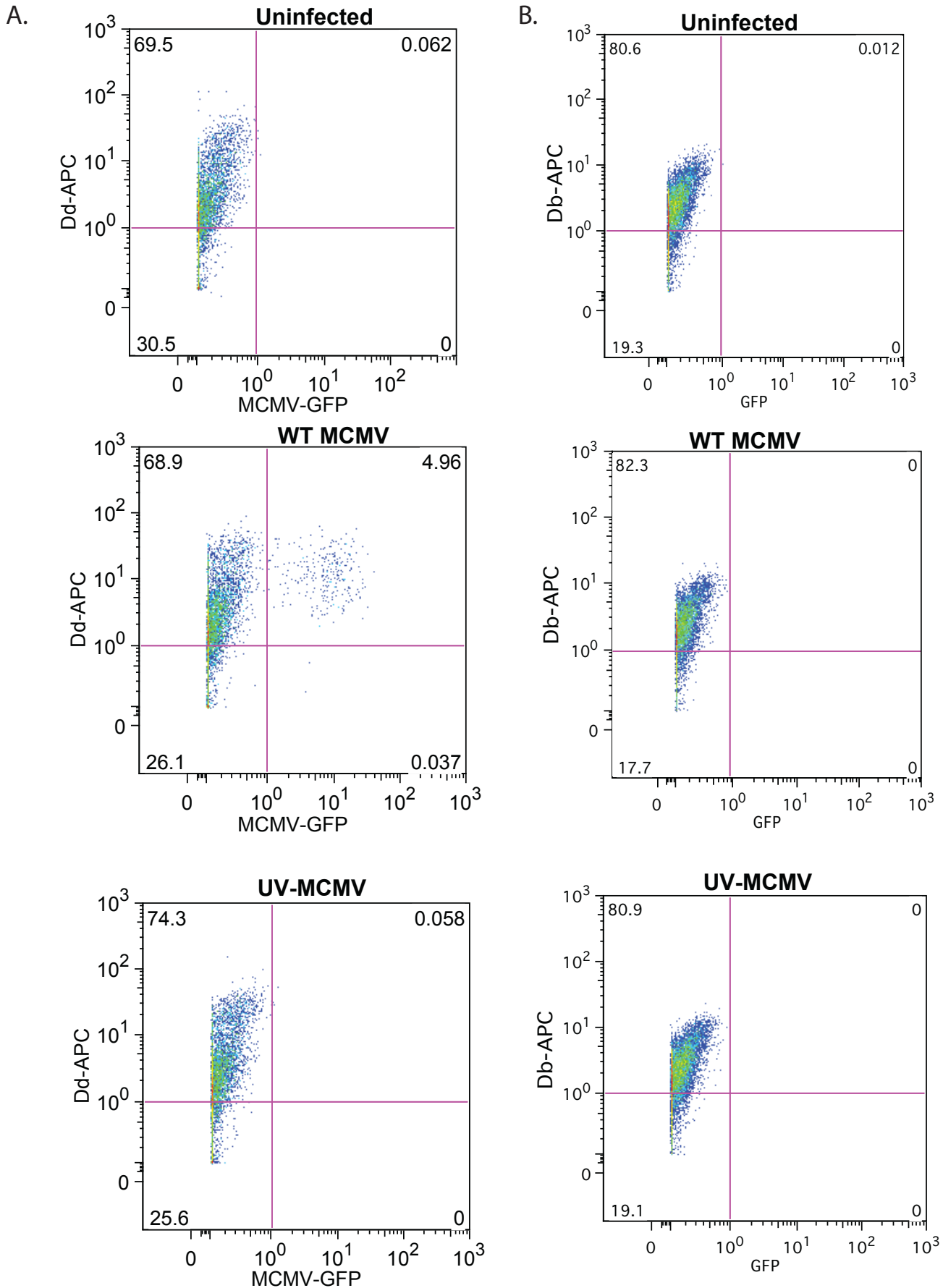


Fig S3: Enhancement of cryptic peptide presentation does not require direct virus infection A. H2d macrophages infected with MCMV-GFP stained with Dd antibody. B. WI9.LYL8 macrophages were stained with a Db antibody and visualized for GFP expression by flow cytometry. Data is representative of 3 experiments.

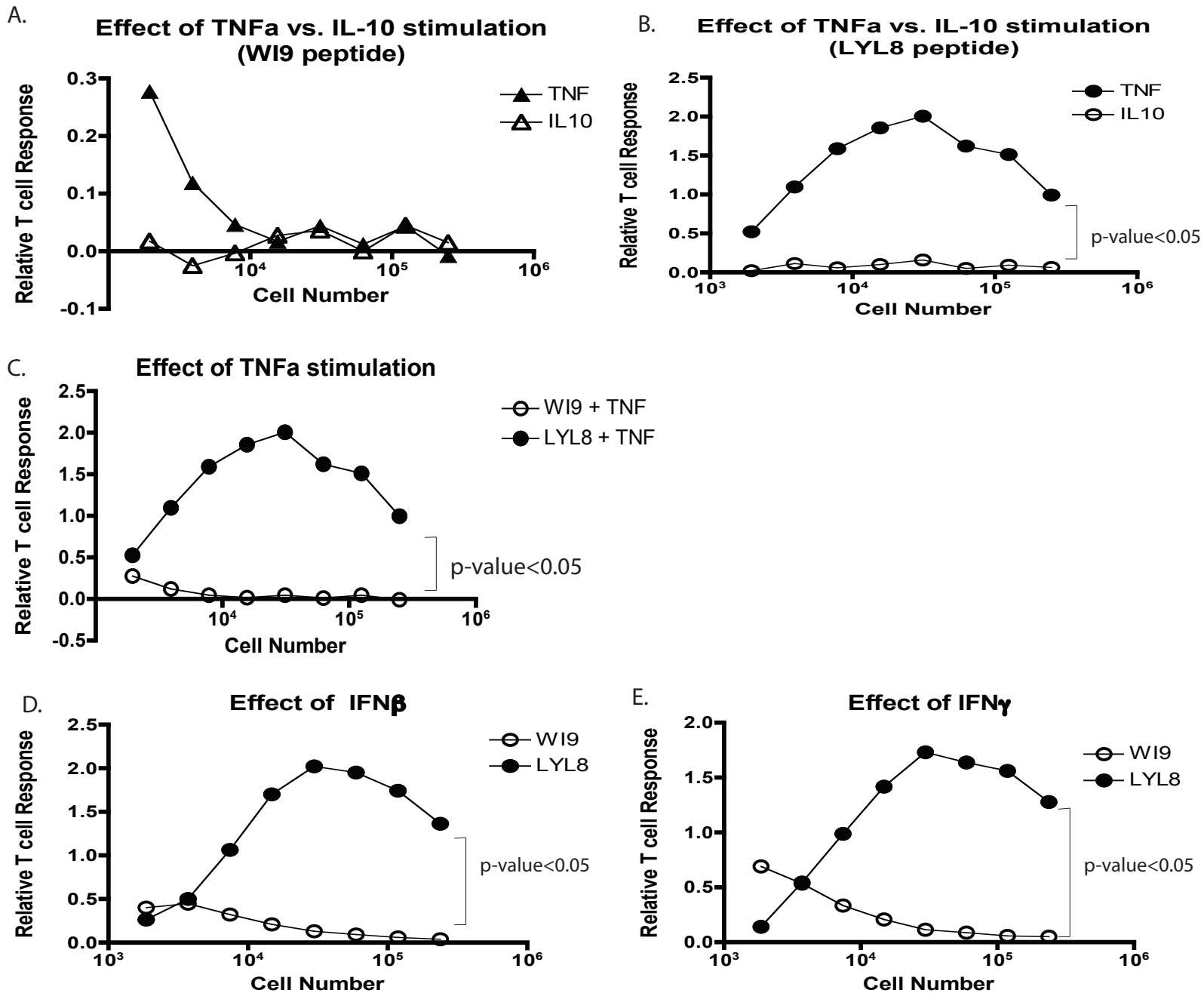


Fig S4 . Relative T-cell responses to the cryptic peptide are significantly enhanced upon inflammatory cytokine stimulation, compared to the conventional peptide. T-cell responses to the WI9 and LYL8 peptides, upon TNF α , IL-10 and IFN β and IFN γ stimulation, were normalized to that of the untreated samples for 3 distinct experiments. An unpaired t-test (with Welch's correction) was performed, comparing the 2 variables. p-value<0.05