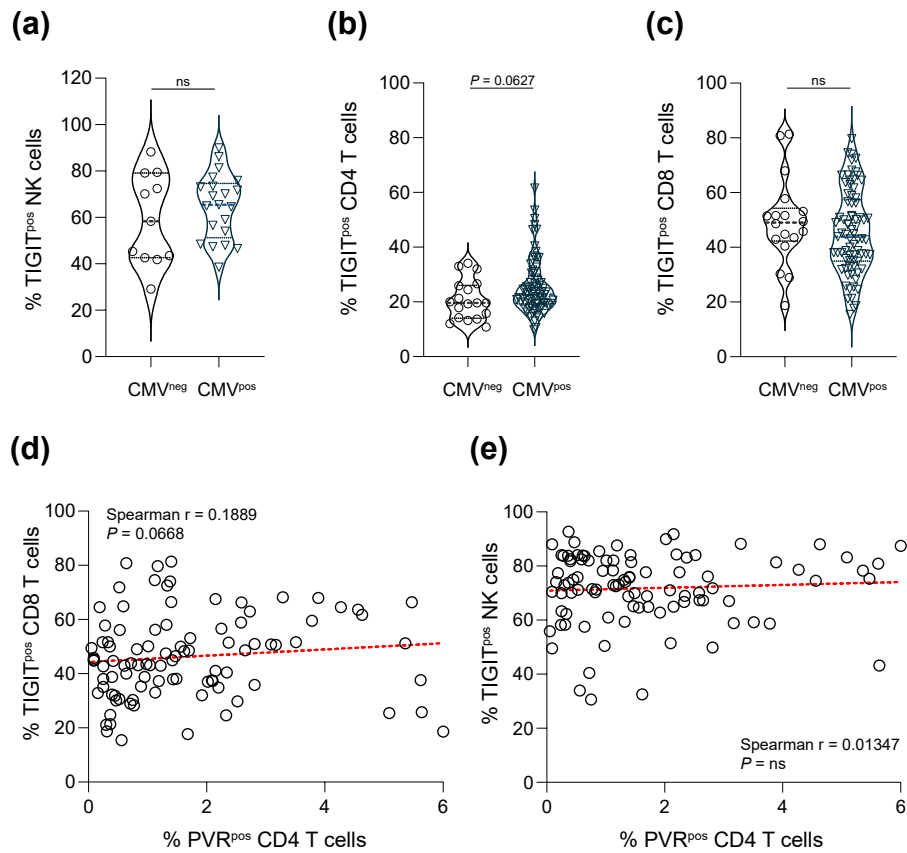
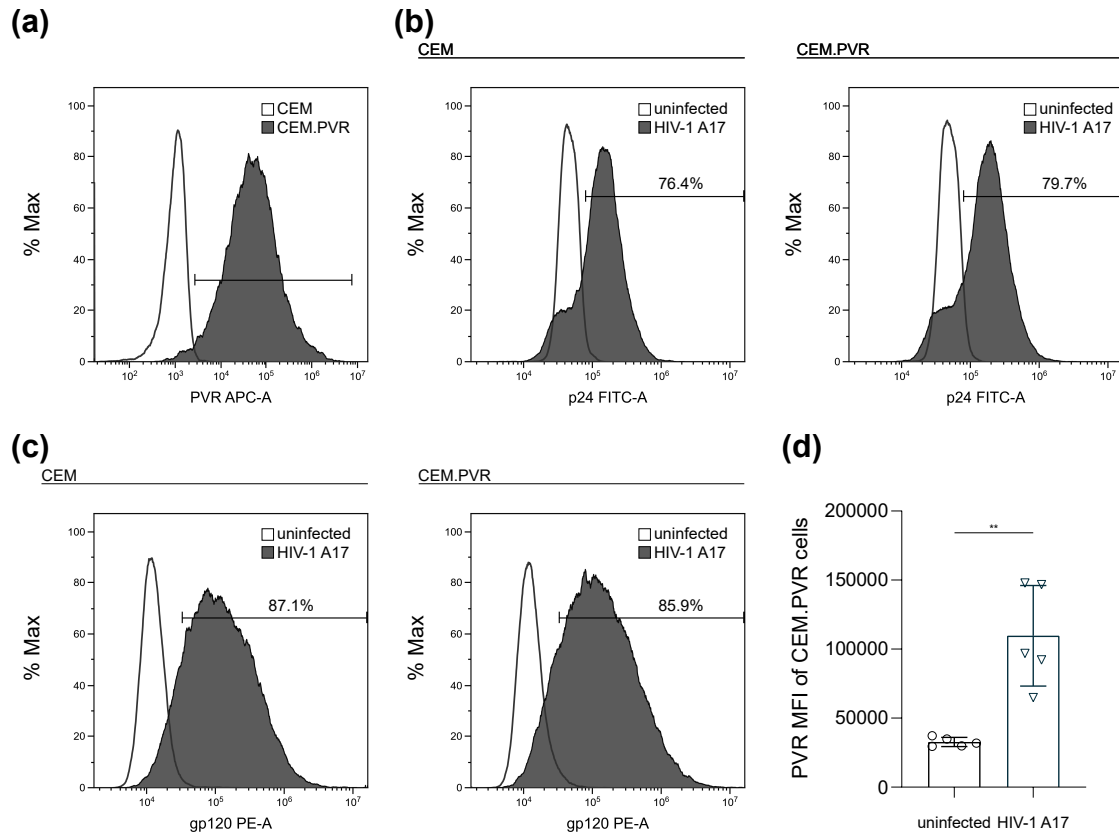


Supplementary Figure 1



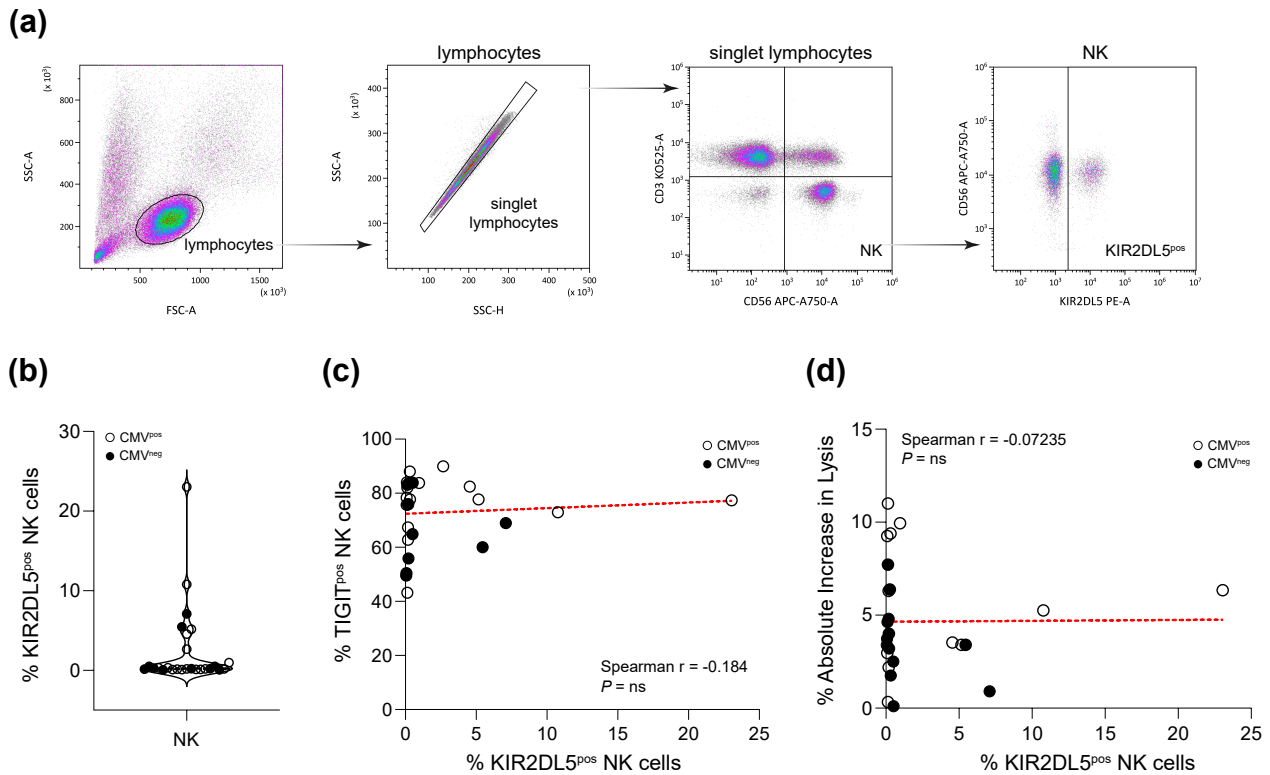
Supplementary Figure 1 | TIGIT expression on lymphocytes from CMV-seronegative and CMV-seropositive participants. (a) Six additional CMV-seronegative individuals were included for more robust comparison of TIGIT expression on NK cells from CMV-seronegative ($n = 11$) and CMV-seropositive ($n = 21$) HIV-seronegative individuals. CMV-seronegative individuals were 64% male with mean age of 49.3 years, whereas CMV-seropositive individuals were 73% male with mean age of 52.0 years. Student's t -test ns. The CMV serostatus of PLWH was determined and (b) CD4⁺ T cell or (c) CD8⁺ T cell TIGIT expression for those who were CMV-seronegative ($n = 18$) was compared to those who were CMV-seropositive ($n = 77$). Mann-Whitney U -test $P = 0.0627$; Student's t -test ns. Correlation between CD4⁺ T cell PVR and percentage of (d) CD8⁺ T cell and (e) NK cell TIGIT expression is depicted. Horizontal lines bisecting groups in (a – c) represent median with IQR.

Supplementary Figure 2



Supplementary Figure 2 | PVR and HIV-1 antigen expression on CEM.NKR-CCR5 cells. Representative histograms from five independent experiments demonstrate **(a)** expression of PVR [median (IQR) 92.3% (5.1%)] on CEM.NKR-CCR5 PVR⁺ (CEM.PVR) cells and relative levels of HIV-1 **(b)** p24 and **(c)** gp120 on CEM.NKR-CCR5 (CEM) and CEM.PVR cells 96 h after HIV-1 A17 infection. The impact of active HIV-1 infection on PVR expression was measured and **(d)** summarized based on MFI of PVR expression (n = 5) on uninfected CEM.PVR cells compared to CEM.PVR cells infected with HIV-1 A17 for 96 h. Student's *t*-test *******P* = 0.0015. Error bars in **(d)** represent SD.

Supplementary Figure 3



Supplementary Figure 3 | Expression of NK cell KIR2DL5 within the functionally studied PLWH cohort. **(a)** NK cells were defined as CD3⁺CD56⁺ singlet lymphocytes and KIR2DL5⁺ NK cells identified. **(b)** Summary data of NK cell KIR2DL5 expression for PLWH within the functionally studied cohort. Correlations between the percentage of NK cells expressing KIR2DL5 and **(c)** TIGIT or **(d)** absolute increase in % specific lysis (raw data from Figure 4c) in the presence of anti-TIGIT mAb vs IgG₁ are depicted. For all graphs, filled symbols represent CMV-seronegative (n = 10) participants while open symbols depict CMV-seropositive participants (n = 16). Horizontal lines bisecting groups in **(b)** represent median with IQR.