

## Figure S1. T cell activation in the CAPRISA cohorts. CD4+ and

CD8+ T cell activation, as measured by the frequency of HLA-DR-positive cells. HIV progressors in the chronic stage of infection (n=18), the two elite controllers (red dots) and HIV-uninfected individuals (n=24) are shown. T cell activation was measured in elite controllers 5 years post HIV infection. Horizontal lines indicate medians. Statistical comparisons were performed using the Mann-Whitney test.



### Figure S2. Flow cytometry plots of HIV-specific CD4+ responses

in elite controller 2 pre- and post-infection. CD4+ T cell IFN-γ, TNF-α and IL-2 cytokine responses from PBMC that were not stimulated (Unstim), or stimulated with Gag or Pol peptides, measured in samples taken 10 months prior to confirmed HIV infection (**A**), 1 month prior to HIV infection (**B**), or 1 month post infection (**C**). Cells were gated on singlets, live CD3+ lymphocytes and then CD4+ cells. The frequency of cytokine positive events is indicated on the plots. A positive cytokine response was defined as at least twice the Unstim sample. Cytokine responses in Figure 1C and F in the manuscript are presented as the background-subtracted total cytokine response, representing the frequency of CD4+ (or CD8+) T cells producing any of the three cytokines. However, due to high TNF-α background responses, TNF-α-single positive cells were excluded from the total response.

		Subtype C						Subtype B						Subtype A						
PID	Years p.i.	DU156.12	DU172.17	CAP45.G3	CAP239.G3	ZM197M.PB7	ZM214M.PL1	6535.3	TRO.11	AC10.0.29	QH0692.42	PVO.4	WITO4160	Q23.17	Q259.d2.17	Q168.a2	Q842.d12	Q769.d22	Q461.e2	% Breadth
EC1	2																			17%
	3																			11%
	4																			6%
EC2	2																			0%
	3																			0%
	4																			44%



#### Figure S3. HIV neutralizing antibody responses in elite

**controllers.** Plasma from two, three and four years post-infection (p.i.) was assayed for neutralization breadth against a panel of 18 heterologous viruses, including 6 subtype C, 6 subtype B and 6 subtype A viruses. Neutralization data are represented in a heat-map, with lack of neutralization indicated in blue,  $IC_{50}$  titers of 1:41-1:100 in yellow, 1:101-1:500 in orange and greater than 1:500 in red. Percentage breadth is indicated.



## Figure S4. Host genetic factor expression in the CAPRISA

**cohorts.** mRNA expression levels for the p21 (**A**) and CCR5 (**B**) genes were measured in HIV progressors (n=10), HIV controllers (n=10) and HIV-uninfected individuals (n=30). ECs are indicated as red dots. Expression levels are represented relative to the housekeeping gene GAPDH. Horizontal lines indicate medians. Statistical comparisons were performed using the Mann-Whitney test.