Figure S1



Figure S1. Controls for removal of HIV-1 specific IgG by Protein G column. 150µl of 10 µg/ml IgG or RPMI was absorbed by Protein G as the experiments in Figure 4. **A.** No detectable HIV-1 specific IgG in the flow-through after Protein G absorption. HIV-1 specific IgG in the flow-through fraction was measured by a HIV-1 multiplex binding assay (HIVIG, 2G12, and G8 against ConSgp140, 4E10 against HIV-1 Env gp41) after passing through the Protein G column. **B.** The flow-through of the 2G12 mAb after absorption by Protein G columns did not inhibit virus replication in a TZM-bl neutralization assay.

Figure S2



Figure S2. Analysis of complete *env* sequences from captured and uncaptured (flow-through) virions. The *env* sequences were obtained by SGA and analyzed using **A**. the Neighbor-Joining tree and **B**. Highlighter plot methods. The sequences from captured viruses are indicated by red color and those from flow-through viruses by blue color. Sequences identical to each other in each phylogenetic lineage are indicated by asterisks. The horizontal scale represents genetic distance.



Figure S3. HIV-1-C3d IC were detectable in plasma from AHI. HIV-1-C3d IC in plasma were captured by mouse anti-C3d IgG coated microplate as described in (<u>43</u>). The captured C3d-virion IC were lysed and measured by viral RNA real time RT-PCR. The HIV-1 –C3d IC was presented as viral RNA copies/ml (Black circle). The percentage of the HIV-1-C3d IC to the total plasma viral load was calculated and shown here (gray circle). The results of 2 typical subjects were shown here.

Figure S4



Figure S4. The endogenous IgG-virion IC plotted vs. the dissociation rates for anti-gp41 IgG in 3 additional AHI subjects.

| Cohort | Subjects ID | Captured Virions (RNA) | Total Virions (RNA)* | Peak of Endogenous IgG virion IC |
|----------------------|-------------|------------------------|----------------------|-------------------------------------|
| | | (copies/ml) | (copies/ml) | (%) |
| Plasma Donor | 9012 | 4554 | 11529 | 39.5 |
| ZeptoMetrix | 9015 | 78000 | 690265 | 11.3 |
| N=13 | 9018 | 27775 | 143170 | 19.4 |
| | 9021 | 219250 | 504023 | 43.5 |
| | 9023 | 36480 | 388085 | 9.4 |
| | 9030 | 123000 | 549107 | 22.4 |
| | 9034 | 31660 | 245426 | 12.9 |
| | 9075 | 119800 | 469804 | 25.5 |
| | 9076 | 1819 | 2584 | 70.4 |
| | 9077 | 3216 | 8787 | 36.6 |
| | 12008 | 3232 | 35130 | 9.2 |
| | 6244 | 4299 | 102357 | 4.2 |
| | 6247 | 342 | 8143 | 4.2 |
| Plasma Donor | PRRG3Q | 687 | 1544 | 44.5 |
| Seracare | DPRG//3 | 2725 | 28380 | 7 1 |
| N=4 | PR8951 | 33275 | 350263 | 9.5 |
| | DDDDCT | 169750 | 1062200 | 9.5 0 C |
| | C1 0275 | 6760 | 75111 | 0.0 |
| 001 | C1=0275 | 0700 | /5111 | 5.0 |
| N=21 | C1-0427 | 1167 | 2785 | 41 9 |
| | C1-0570 | 0 | 7043 | 0 |
| | C1-0666 | 9360 | 95510 | 9.8 |
| | C1-0709 | 1212 | 7528 | 16.1 |
| | C1 09E0 | 20040 | 226525 | 10.1 |
| | C1 0259 | 70800 | 128000 | 12.7 62.2 |
| | C1 0517 | 79800 | 122030 | 69.2 |
| | C1-0317 | 30800 | 132545 | 08.5 |
| | C1-0149 | 1856 | 4052 | 45.8 |
| | C1-0185 | 3868 | 1//43 | 21.8 |
| | C1-0047 | 8700 | 20519 | 42.4 |
| | C1-0157 | 1112 | 5032 | 22.1 |
| | C1-0736 | 4632 | 11847 | 39.1 |
| | C1-0798 | 170400 | 230582 | 73.9 |
| | C1-0455 | 55160 | 100109 | 55.1 |
| | C1-0976 | 9400 | 17871 | 52.6 |
| | C1-1023 | 8833 | 22138 | 39.9 |
| | C1-1096 | 37120 | 74240 | 50.0 |
| | C1-1215 | 19080 | 61154 | 31.2 |
| | C1-0731 | 5236 | 16061 | 32.6 |
| | C1-0010 | 32229 | 51649 | 62.4 |
| Chronic CHAVI 001 | C1-0586 | 19600 | 29297 | 66.9 |
| N=10 | C1-0645 | 75720 | 128122 | 59.1 |
| | C1-0237 | 940 | 2892 | 32.5 |
| | C1-0060 | 407 | 651 | 62.5 |
| | C1-0642 | 18520 | 26647 | 69.5 |
| | C1-0588 | 19060 | 41435 | 46.0 |
| | C1-0261 | 5526 | 8771 | 63.0 |
| | C1-0300 | 2904 | 6785 | 42.8 |
| | C1-0573 | 177 | 276 | 64 1 |
| | C1-0502 | 2082 | 4090 | 5 <u>.</u> |

Table S1

*Total virions (RNA) = the viral RNA of the flow-through + the viral RNA of capture in Protein G capture assay

Table S1. Summary of the peak of plasma endogenousIgG coated virion IC in 38 AHI and 10 chronic subjects.

Table S2

| | f = 1 | f = 14 | f = 35 |
|---------|--|--|--|
| Patient | $K \; (l/mol)$ | K (l/mol) | $K \; (l/mol)$ |
| PRB951 | 7.9×10^5 $[0, 5.8 \times 10^6]$ | 5.4×10^4 $[0, 3.2 \times 10^5]$ | 2.2×10^4 [0, 1.2 × 10 ⁵] |
| PRB957 | 6.2×10^4 [0, 4.6 × 10 ⁵] | 4.3×10^3 [0, 2.5 × 10 ⁴] | 1.7×10^3 $[0, 9.8 \times 10^3]$ |
| PRB939 | $4.2 \times 10^4 [1.9 \times 10^4, 8.4 \times 10^4]$ | $2.2 \times 10^{3} \\ [1.1 \times 10^{3}, 3.7 \times 10^{3}]$ | $8.8 \times 10^2 [4.7 \times 10^2, 1.4 \times 10^3]$ |
| 9030 | 5.2×10^{6} $[2.4 \times 10^{6}, 8.8 \times 10^{6}]$ | $\begin{array}{c} 3.3 \times 10^5 \\ [1.5 \times 10^5, 5.3 \times 10^5] \end{array}$ | $\frac{1.3 \times 10^5}{[6 \times 10^4, 2.1 \times 10^5]}$ |
| 9075 | $\frac{3.7 \times 10^4}{[7.7 \times 10^3, 8.6 \times 10^4]}$ | $2.2 \times 10^3 [4.2 \times 10^2, 4.7 \times 10^3]$ | 8.9×10^{2} $[1.6 \times 10^{2}, 1.9 \times 10^{3}]$ |
| 9012 | $\frac{4.7 \times 10^6}{[2.7 \times 10^6, 7.8 \times 10^6]}$ | $\frac{2.6 \times 10^5}{[1.6 \times 10^5, 3.9 \times 10^5]}$ | $\frac{1 \times 10^5}{[6.4 \times 10^4, 1.5 \times 10^5]}$ |
| 9034 | $3.1 	imes 10^6 \ [0, 9.6 	imes 10^6]$ | 2.1×10^5 [0, 6.2 × 10 ⁵] | 8.5×10^4 [0, 2.4 × 10 ⁵] |
| 12008 | $\frac{8 \times 10^4}{[7.7 \times 10^3, 1.7 \times 10^5]}$ | $5.4 \times 10^{3} \\ [5 \times 10^{2}, 1.1 \times 10^{4}]$ | $2.2 \times 10^{3} \\ [2 \times 10^{2}, 4.3 \times 10^{3}]$ |

Table S2. The estimated affinity constant K for each patient given a model assuming either univalent (f=1) or multivalent binding with f=14 or f=35 maximal number of possible binding sites. Numbers in brackets represent 95% confidence intervals of the estimates.