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## Supplementary Information for

HLA tapasin independence: broader peptide repertoire and HIV control

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## **Supplementary Information Text**

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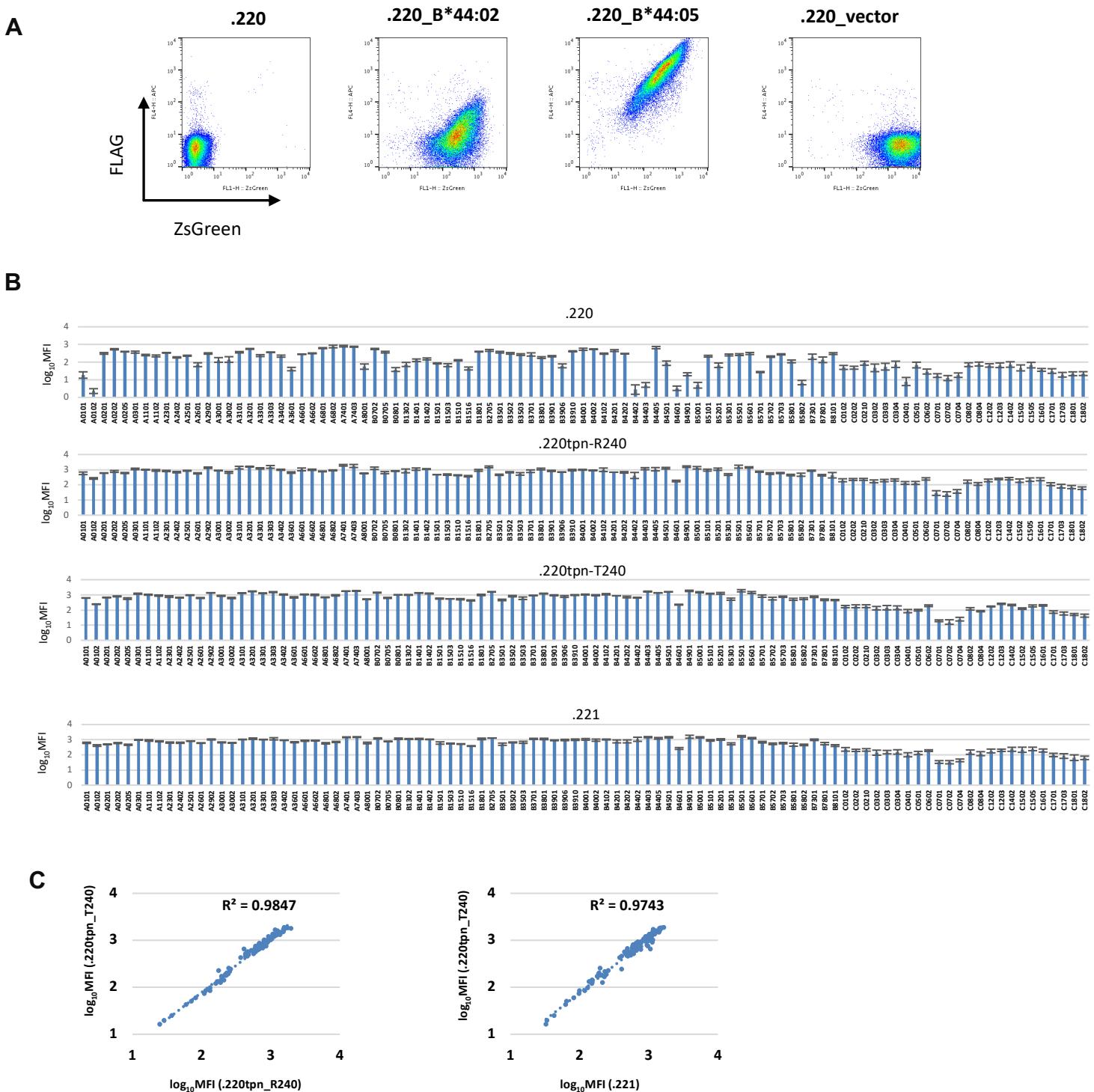
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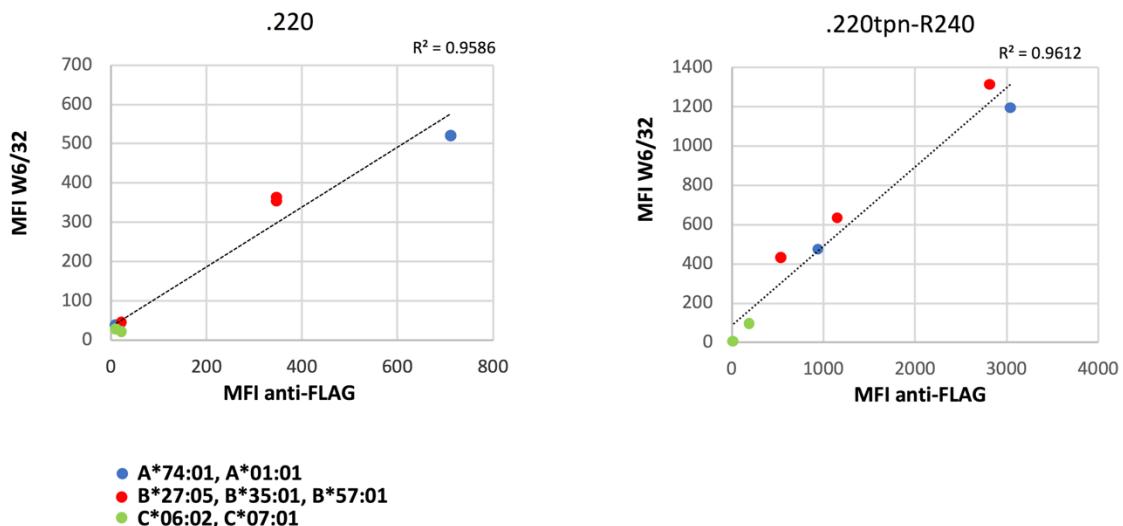
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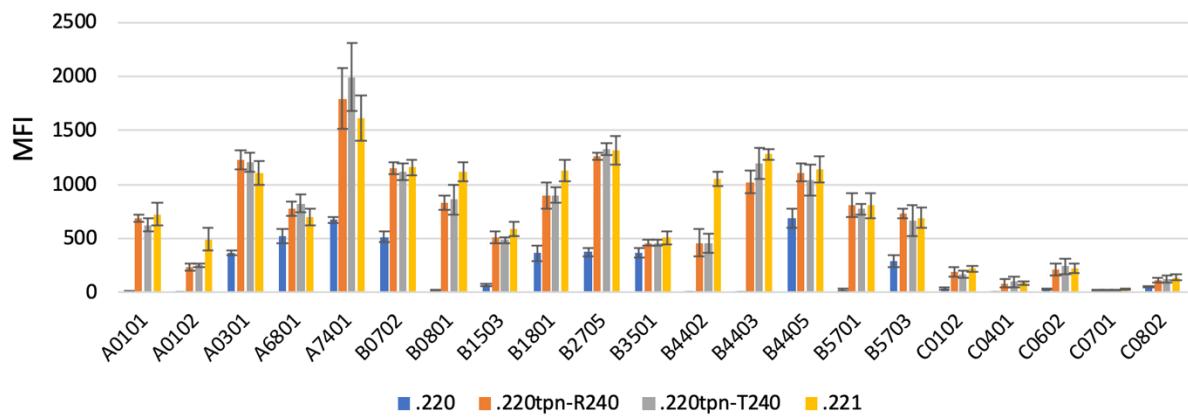
the Swiss HIV Cohort Study: Aebi-Popp K, Anagnostopoulos A, Battegay M, Bernasconi E, Böni J, Braun DL, Bucher HC, Calmy A, Cavassini M, Ciuffi A, Dollenmaier G, Egger M, Elzi L, Fehr J, Fellay J, Furrer H, Fux CA, Günthard HF (President of the SHCS), Haerry D (deputy of "Positive Council"), Hasse B, Hirsch HH, Hoffmann M, Hösli I, Huber M, Kahlert CR (Chairman of the Mother & Child Substudy), Kaiser L, Keiser O, Klimkait T, Kouyos RD, Kovari H, Ledergerber B, Martinetti G, Martinez de Tejada B, Marzolini C, Metzner KJ, Müller N, Nicca D, Paioni P, Pantaleo G, Perreau M, Rauch A (Chairman of the Scientific Board), Rudin C, Scherrer AU (Head of Data Centre), Schmid P, Speck R, Stöckle M (Chairman of the Clinical and Laboratory Committee), Tarr P, Trkola A, Vernazza P, Wandeler G, Weber R, Yerly S.



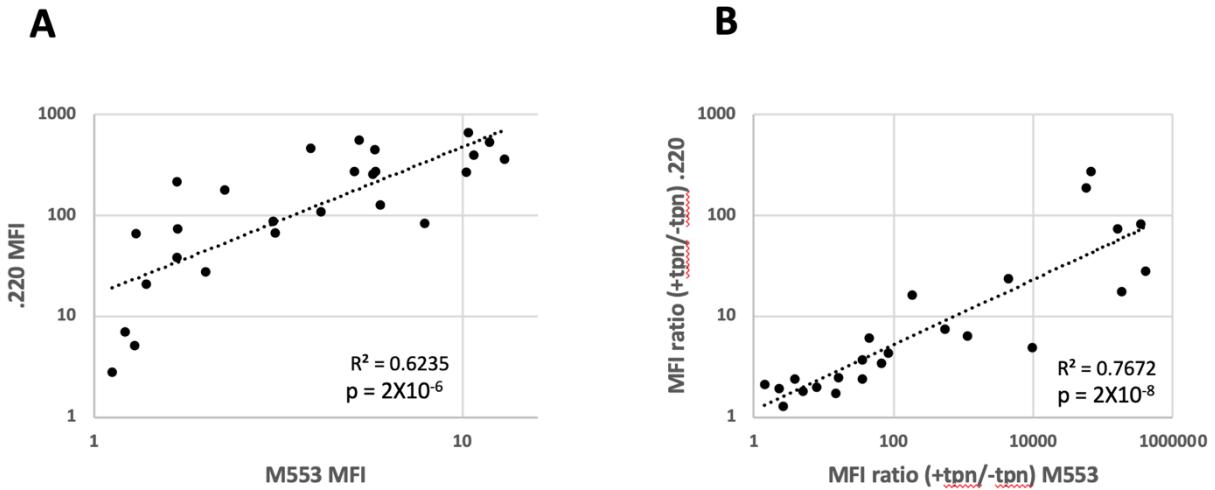
**Fig. S1. Flow cytometric analysis of HLA expression in the presence or absence of tapasin.** (A) Representative flow cytometry plots show ZsGreen fluorescence and anti-FLAG mAb binding to parental .220 cells, and .220 cells transduced with *HLA-B\*44:02*, *HLA-B\*44:05*, or “empty vector” (no HLA) constructs. ZsGreen fluorescence marks cells that were lentivirally transduced. Tapasin-dependent *HLA-B\*44:02* is expressed at lower levels compared to tapasin-independent *HLA-B\*44:05*. (B) HLA class I allotype expression levels in .220, .220tpn-R240, .220tpn-T240, and .221 cells. The MFI values and standard deviations represent data obtained from four measurements performed on different days on the same cells maintained in culture. (C) Expression levels measured in .220tpn\_T240 and .220tpn\_R240 cells (left), as well as in .220tpn\_T240 and .221 cells (right) demonstrate significant correlations, respectively.



**Fig. S2. HLA expression levels measured using W6/32 mAb and anti-FLAG mAb strongly correlate.** Expression of two HLA-A, three HLA-B and two HLA-C allotypes were analyzed by flow cytometry on the surface of .220 (left) and .220tpn-R240 (right) cells.



**Fig.S3. Comparison of HLA allotype expression levels across .220, .220tpn, and .221 cells.** Tapasin-negative and -positive cells transduced with a subset of *HLA* alleles were analyzed by flow cytometry using anti-FLAG mAb on the same day. Expression levels in tapasin-positive cells are similar across most allotypes, although A\*01:02, B\*44:02, and, to a lesser extent, B\*08:01 and C\*07:01 showed lower expression on .220tpn than on .221 cells ( $p=0.03$ , Wilcoxon test). These differences may reflect some level of insufficiency of the exogenous compared to endogenous tapasin for assembly of these particular allotypes. There was no difference in expression level of each allele on .220tpn cells transduced with either R240 or T240 tapasin ( $p>0.05$ , Wilcoxon test). The MFI values and standard deviations represent data obtained from four measurements performed on different days using the same cells maintained in culture.



**Fig. S4. Correlation between HLA-B expression on .220 and M553 cells (M553 data derived from Rizvi et al. (14)).** (A) Comparison of HLA-B expression levels in .220 vs. M553 cells. (B) Comparison of HLA-B MFI ratios (+tpn/-tpn) in .220 vs. M553 cells. Each dot represents an HLA-B allotype that was evaluated for TD in both the present study and Rizvi et al.

**Table S1. TD of HLA class I allotypes defined as the ratio of cell surface expression levels in .220tpn over that in .220 cells.**

| HLA     | MFI ratio (+tpn/-tpn) | SD    | HLA     | MFI ratio (+tpn/-tpn) | SD     |
|---------|-----------------------|-------|---------|-----------------------|--------|
| A*01:01 | 32.11                 | 13.77 | B*40:02 | 1.71                  | 0.11   |
| A*01:02 | 109.86                | 35.04 | B*41:02 | 3.45                  | 0.53   |
| A*02:01 | 2.02                  | 0.22  | B*42:01 | 1.72                  | 0.21   |
| A*02:02 | 1.45                  | 0.16  | B*42:02 | 2.31                  | 0.21   |
| A*02:05 | 1.49                  | 0.12  | B*44:02 | 190.40                | 121.20 |
| A*03:01 | 3.14                  | 0.46  | B*44:03 | 274.16                | 85.74  |
| A*11:01 | 4.00                  | 0.40  | B*44:05 | 1.82                  | 0.33   |
| A*11:02 | 3.99                  | 0.57  | B*45:01 | 16.31                 | 4.70   |
| A*23:01 | 2.36                  | 0.20  | B*46:01 | 60.66                 | 17.61  |
| A*24:02 | 3.61                  | 0.35  | B*49:01 | 82.40                 | 16.89  |
| A*25:01 | 3.91                  | 0.29  | B*50:01 | 286.17                | 110.15 |
| A*26:01 | 8.33                  | 2.14  | B*51:01 | 4.92                  | 0.66   |
| A*29:02 | 4.38                  | 0.45  | B*52:01 | 17.51                 | 5.42   |
| A*30:01 | 6.52                  | 2.01  | B*53:01 | 1.96                  | 0.31   |
| A*30:02 | 4.54                  | 1.71  | B*55:01 | 6.46                  | 1.15   |
| A*31:01 | 3.67                  | 0.50  | B*56:01 | 4.60                  | 0.73   |
| A*32:01 | 2.92                  | 0.20  | B*57:01 | 28.43                 | 3.42   |
| A*33:01 | 5.34                  | 0.71  | B*57:02 | 2.74                  | 0.41   |
| A*33:03 | 4.20                  | 0.48  | B*57:03 | 2.46                  | 0.24   |
| A*34:02 | 4.66                  | 0.67  | B*58:01 | 4.30                  | 0.86   |
| A*36:01 | 16.33                 | 3.80  | B*58:02 | 73.28                 | 22.75  |
| A*66:01 | 3.77                  | 0.42  | B*73:01 | 3.97                  | 1.38   |
| A*66:02 | 3.16                  | 0.27  | B*78:01 | 3.33                  | 1.23   |
| A*68:01 | 1.17                  | 0.09  | B*81:01 | 1.45                  | 0.34   |
| A*68:02 | 1.19                  | 0.21  | C*01:02 | 3.67                  | 1.09   |
| A*74:01 | 2.31                  | 0.28  | C*02:02 | 4.17                  | 0.95   |
| A*74:03 | 2.40                  | 0.31  | C*02:10 | 2.34                  | 0.67   |
| A*80:01 | 9.70                  | 3.34  | C*03:02 | 3.19                  | 1.65   |
| B*07:02 | 2.37                  | 0.28  | C*03:03 | 3.01                  | 1.42   |
| B*07:05 | 1.72                  | 0.24  | C*03:04 | 2.32                  | 1.01   |
| B*08:01 | 23.78                 | 5.94  | C*04:01 | 13.59                 | 7.47   |
| B*13:02 | 12.35                 | 3.68  | C*05:01 | 1.68                  | 0.66   |
| B*14:01 | 9.48                  | 1.76  | C*06:02 | 7.43                  | 2.52   |
| B*14:02 | 7.68                  | 1.01  | C*07:01 | 1.38                  | 0.45   |
| B*15:01 | 6.10                  | 0.49  | C*07:02 | 1.63                  | 0.66   |
| B*15:03 | 7.52                  | 1.49  | C*07:04 | 1.69                  | 0.59   |
| B*15:10 | 3.72                  | 0.36  | C*08:02 | 1.97                  | 0.57   |
| B*15:16 | 9.18                  | 1.82  | C*08:04 | 1.28                  | 0.36   |
| B*18:01 | 2.40                  | 0.23  | C*12:02 | 2.83                  | 0.69   |
| B*27:05 | 3.38                  | 0.47  | C*12:03 | 3.76                  | 1.17   |
| B*35:01 | 1.26                  | 0.16  | C*14:02 | 3.19                  | 1.19   |
| B*35:02 | 2.38                  | 0.29  | C*15:02 | 3.31                  | 1.39   |
| B*35:03 | 2.10                  | 0.36  | C*15:05 | 2.99                  | 1.17   |
| B*37:01 | 3.13                  | 0.72  | C*16:01 | 5.98                  | 1.33   |
| B*38:01 | 6.39                  | 0.84  | C*17:01 | 2.92                  | 0.98   |
| B*39:01 | 4.10                  | 0.41  | C*17:03 | 3.63                  | 1.29   |
| B*39:06 | 11.95                 | 3.11  | C*18:01 | 2.76                  | 0.80   |
| B*39:10 | 2.32                  | 0.21  | C*18:02 | 2.28                  | 0.66   |
| B*40:01 | 1.90                  | 0.28  |         |                       |        |