**Table S1: Protease inhibitor resistance mutations and relative CARD8 cleavage.**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **Clone name** | **Reported PI-resistance  mutations in HIVPR** | **HIVgag mutation** | **Strongest**  **PI-R1** | **% CARD8 cleavage**  **(relative to HIVLAI)** | **Additional amino acid changes in HIVPR relative to NL4.3** |
| HIV-1LAI | wildtype | wildtype | wildtype | 100 | 37S |
| CA126802  (PI-R1) | 11I, 32I, 33F, 46I, 47V, 54M, 58E, 73S, 84V, 89V, 90M | 431V (NC/p1)  437N (NC/p1)  453LF (p1/p6) | FPV, LPV, TPV, DRV | 51 | 10I, 12K, 13V, 20V, 35G, 36I, 37D, 57K, 63P, 64V, 66V, 7IV |
| **CA122805 (PI-R2)** | **10F, 33F, 43T, 46L, 54V, 82A, 84V, 90M** | **431V (NC/p1)**  **532S (p1/p6)** | **SQV** | **7** | 16A,19I, 20R, 35D, 36L, 37D, 55R, 57K, 60E, 62V, 63P, 71V, 93L |
| CA126805 (PI-R3) | 33F, 43T, 46I, 48V, 50V, 54S, 82A | 437N (NC/p1)  449VF (p1/p6)  PTAP insertion (p6) | SQV, IDV, LPV | **8** | not determined\* |
| CA96457  (PI-R5) | 48V, 53L, 54V, 82A, 90M | 436R  (NC/p1) | SQV | 11 | 10I, 37D, 63P, 71T, 77I, 93L |
| **CA96458**  **(PI-R9)** | **10F, 30N, 33F, 43T, 84V, 88D, 90M** | **431V (NC/p1)** | **NFV, SQV** | **24** | 15V, 35D, 36L, 37E, 60E, 62V, 63P |
| CA50834-1  (PI-R10) | 24I, 46L, 54V, 76V, 82A | 431V (NC/p1) | IDV, LPV | 38 | 10I, 14R, 35D, 36I, 37E, 63P,71V |
| **CA96451**  **(PI-R12)** | **32I, 33F, 43T, 46I, 47V, 54M, 73S, 82A, 89V, 90M** | **437N (NC/p1)**  **PTAP insertion (p6)** | **FPV, LPV** | **192** | 10V, 12V, 13V, 15V, 20M, 60E, 61N, 62V, 63P, 67Y, 69K, 71I, 72L, 77I |
| **CA20392-1**  **(PI-R13)** | **24I, 46L, 54V, 82A** | **431V (NC/p1)** | **LPV** | **206** | 10I, 14R, 35D, 36I, 37E, 63P, 71V |