

PI Major Mutations:

M46M

PI Accessory Mutations:

None

PR Other Mutations:

I13V • Q18L • L19LQ • L33V • M36I • R41K • L63P • I64V • I72V

Protease Inhibitors	
atazanavir/r (ATV/r)	Potential Low-Level Resistance
darunavir/r (DRV/r)	Susceptible
lopinavir/r (LPV/r)	Potential Low-Level Resistance

PR comments

Major

- M46I/L are relatively non-polymorphic PI-selected mutations. In combination with other PI-resistance mutations, they are associated with reduced susceptibility to each of the PIs except DRV.

Other

- L33I/V are minimally polymorphic mutations that do not appear to be selected by PIs or to reduce their susceptibility.

Drug resistance mutation scores of PI:

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Rule	ATV/r	DRV/r	LPV/r
M46M	10	0	10

NRTI Mutations:

K70Q • M184I

NNRTI Mutations:

K101E • K103N • G190A

RT Other Mutations:

K11KR • V33I • T39EK • E40EK • M41M • E42EK • V60I • R72RK • V90I • S105T • D121Y • K122E • I135K • V148VM • Q174QR • D177E • I178MV • E194EK • T200I • E204EK • Q207E • R211K • L228R • V245T • D250E • G262GE • A272P • G273GE • K275KR • L282C • L283I • R284RK • G283K • T286A • A288ST • I293V

Nucleoside Reverse Transcriptase Inhibitors		Non-nucleoside Reverse Transcriptase Inhibitors	
abacavir (ABC)	Intermediate Resistance	doravirine (DOR)	Low-Level Resistance
zidovudine (AZT)	Susceptible	efavirenz (EFV)	High-Level Resistance
emtricitabine (FTC)	High-Level Resistance	etravirine (ETR)	Intermediate Resistance
lamivudine (3TC)	High-Level Resistance	nevirapine (NVP)	High-Level Resistance
tenofovir (TDF)	Low-Level Resistance	rilpivirine (RPV)	High-Level Resistance

RT comments

NRTI

- K70/E/Q/N/T/S/G cause low-leve resistance to ABC and TDF.
- M184V/I cause high-level in vitro resistance to 3TC and FTC and low/intermediate resistance to ABC (3-fold reduced susceptibility). M184V/I are not contraindications to continued treatment with 3TC or FTC because they increase susceptibility to AZT and TDF and are associated with clinically significant reductions in HIV-1 replication.

NNRTI

- K101E is a non-polymorphic accessory mutation that confers intermediate resistance to NVP and RPV and low-level reductions in susceptibility to EFV, ETR, and DOR when it occurs with other NNRTI-resistance mutations.
- K103N is a non-polymorphic mutation that confers high-level reductions in NVP and EPV susceptibility. It is the most commonly transmitted DRM.
- G190A is a non-polymorphic mutation that causes high-level resistance to NVP and intermediate resistance to EFV. It does not significantly reduce susceptibility to RPV, ETR, or DOR.

Other

- V90I is a polymorphic accessory mutation weakly selected by each of the NNRTIs. It is associated with minimal, if any, detectable reduction in NNRTI susceptibility.

Drug resistance mutation scores of NRTI:

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Rule	ABC ⚡	AZT ⚡	FTC ⚡	3TC ⚡	TDF ⚡
<u>K70Q</u>	15	0	10	10	15
<u>M184I</u>	15	-10	60	60	-10
<u>K70Q + M184I</u>	0	0	0	0	10
Total	30	-10	70	70	15

Drug resistance mutation scores of NNRTI:

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Rule	DOR ⚡	EFV ⚡	ETR ⚡	NVP ⚡	RPV ⚡
<u>K101E</u>	15	15	15	30	45
<u>K101E + G190A</u>	5	0	5	0	0
<u>K103N</u>	0	60	0	60	0
<u>G190A</u>	0	45	10	60	15
<u>K101E + M184I</u>	0	0	0	0	15
Total	20	120	30	150	75