HIVDB 9.5.1 (2023-11-05) Drug resistance interpretation: PR

PI Major Mutations: None PI Accessory Mutations: None

V11X - T12A - I13S - K14\* - I15D - G16R - Q18K - K20R - E35D - M36I - N37K - R41K - R57K - H69K - K70R - L89M PR Other Mutations:

# Protease Inhibitors

atazanavir/r (ATV/r) Susceptible Susceptible darunavir/r (DRV/r) Susceptible fosamprenavir/r (FPV/r) Susceptible indinavir/r (IDV/r) lopinavir/r (LPV/r) Susceptible nelfinavir (NFV) Susceptible saguinavir/r (SQV/r) Susceptible Susceptible tipranavir/r (TPV/r)

#### PR comments

tenofovir (TDF)

### Other

K20R is a highly polymorphic PI-selected accessory mutation that increases replication fitness in viruses with PI-resistance mutations.

HIVDB 9.5.1 (2023-11-05) Mutation scoring: PR

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No drug resistance mutations were found for PI.

# Drug resistance interpretation: RT

NRTI Mutations: K65R - S68G - M184V NNRTI Mutations: L1001 - K103N - E138G

E6D - K20R - V35T - E40D - K49R - V60I - K122E - D123S - I135T - F171X - K173S - Q174R - V179I - Q182X - T200A - I202V - E203Q - Q207D - E224D - Y232M - E233N - A234 - H235S - P236S - P243L - I244S - V245E - P247X - K249E - N255M - D256I - I257Y - Q258R - K259I - L260V - V261E - G262T -RT Other Mutations:

L264W • N265Q • W266S • A267I • S268M • Δ270-273 • I274X • K277R • K281S • L282S • L283S • R284A • G285D

# **Nucleoside Reverse Transcriptase Inhibitors**

# Non-nucleoside Reverse Transcriptase Inhibitors

abacavir (ABC) High-Level Resistance doravirine (DOR) Intermediate Resistance efavirenz (EFV) High-Level Resistance zidovudine (AZT) Susceptible stavudine (D4T) Intermediate Resistance etravirine (ETR) Intermediate Resistance nevirapine (NVP) didanosine (DDI) High-Level Resistance High-Level Resistance emtricitabine (FTC) High-Level Resistance rilpivirine (RPV) High-Level Resistance High-Level Resistance lamivudine (3TC) Intermediate Resistance

### RT comments

### NRTI

- K65R confers intermediate reductions in susceptibility to TDF, ABC, and 3TC/FTC. It increases AZT susceptibility. In NRTI-experienced, INSTI-naive patients receiving TDF+3TC+DTG, there is a risk of emergent DTG resistance that does not arise in NRTI-naive patients receiving TDF+3TC+DTG.
- \$686 is a polymorphic mutation that is often selected in combination with K65R. It partially restores the replication defect associated with K65R.
- 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

- L100I is a non-polymorphic mutation that usually occurs in combination with K103N. In this setting it confers high-level resistance to NVP, EFV, and RPV and intermediate resistance to ETR and DOR.
- K103N is a non-polymorphic mutation that confers high-level reductions in NVP and EFV susceptibility. It is the most commonly transmitted DRM.
- . E138Q/G are non-polymorphic accessory mutations selected by ETR occasionally NVP and EFV. They cause low-level reductions in susceptibility to NVP, RPV, and ETR.

# Other

- V179I is a polymorphic mutation that is frequently selected in persons receiving ETR and RPV. However, it has little, if any, direct effect on NNRTI susceptibility.
- L234I is a nonpolymorphic mutation selected in persons receiving NVP and EFV. It is also selected in vitro by ETR and DOR. In combination with V106A, it is associated with high-level DOR resistance. Its effect on susceptibility when it occurs alone has not been well characterized. L234del is a highly unusual mutation at this position.
- . P236L is a rare mutation selected commonly by DLV, which appears to have little if any effect on current NNRTIs. P236S is a highly unusual mutation at this position.

HIVDB 9.5.1 (2023-11-05) Mutation scoring: RT

Drug resistance mutation scores of NRTI:



Rule	ABC ≑	AZT ≑	D4T ≑	DDI 🗦	FTC ‡	зтс ≑	TDF ‡
K65R	45	-10	60	60	30	30	50
M184V	15	-10	-10	10	60	60	-10
K65R + S68G	0	0	0	0	0	0	5
Total	60	-20	50	70	90	90	45

Drug resistance mutation scores of NNRTI:



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Rule	DOR \$	EFV \$	ETR \$	NVP ÷	RPV \$
L100I	15	60	30	60	60
L100I + K103N	15	0	0	0	0
K103N	0	60	0	60	0
E138G	0	10	10	10	15
Total	30	130	40	130	75