PI Major Mutations: None PI Accessory Mutations: None PR Other Mutations: Protease Inhibitors

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Drug resistance interpretation: PR

Susceptible atazanavir/r (ATV/r) darunavir/r (DRV/r) Susceptible Susceptible lopinavir/r (LPV/r)

PR comments

Mutation scoring: PR

Other

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

Drug resistance interpretation: RT

No drug resistance mutations were found for Pl.

NRTI Mutations: D67N 94% K70E 95% M184IV V: 18%, E 18%

V179D 52% - Y188L 95% com 34 572

NNRTI Mutations: RT Other Mutations:

E6D 99% - V35T 99% - T39E 99% - V245Q 99% - D250E 97% - D250E 97%

G335D 976 - R356K 986 - M357R 946 - G359T 976 - K366R 976

Nucleoside Reverse Transcriptase Inhibitors Non-nucleoside Reverse Transcriptase Inhibitors Intermediate Resistance doravirine (DOR) High-Level Resistance

abacavir (ABC) High-Level Resistance zidovudine (AZT) Susceptible efavirenz (EFV) emtricitabine (FTC) High-Level Resistance etravirine (ETR) Low-Level Resistance lamivudine (3TC) High-Level Resistance nevirapine (NVP) High-Level Resistance tenofovir (TDF) Low-Level Resistance rilpivirine (RPV) High-Level Resistance

RT comments

NRTI

- . D67N is a non-polymorphic TAM associated with low-level resistance to AZT.
- 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

- V179D/E are somewhat polymorphic accessory NNRTI-selected mutation. In combination with other NNRTI DRMs, they appear to contribute low-levels of reduced susceptibility to each of the NNRTIs. In particular, the combinations of K103R/V179D and V106I/V179D act synergistically to reduce NVP and EFV susceptibility.
- Y188L is a non-polymorphic mutation that confers high-level resistance to NVP, EFV, RPV, and DOR, and potentially low-level resistance to ETR.

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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.
- . K103R is a polymorphic mutation that alone has no effect on NNRTI susceptibility. However, in combination with V179D, it reduces NVP and EFV susceptibility about 15-fold.

Mutation scoring: RT

Drug resistance mutation scores of NRTI:

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Rule	ABC ÷	AZT \$	FTC ÷	зтс ≑	TDF \$
<u>D67N</u>	5	15	0	0	5
<u>K70E</u>	15	0	10	10	15
M184IV	15	-10	60	60	-10
K70E + M184IV	0	0	0	0	10
Total	35	5	70	70	20

Drug resistance mutation scores of NNRTI:

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Rule	DOR \$	EFV ≑	ETR ≑	NVP ≑	RPV \$
<u>Y188L</u>	60	60	10	60	60
K103KR + V179D	0	20	0	20	15
<u>V179D</u>	0	10	10	10	10
Total	60	90	20	90	85