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

PI Major Mutations: None PI Accessory Mutations: None

PR Other Mutations: 113V Q18E 135D N36I R41K K45R K45R 163K L63A H69K L89M

### Protease Inhibitors

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

#### Mutation scoring: PR

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No drug resistance mutations were found for PI.

Drug resistance interpretation: RT

NRTI Mutations: D67DN o see a K70R see M184V mg T215F mg K219E see

NNRTI Mutations: K101H --- V108W --- G190A ---

#### Nucleoside Reverse Transcriptase Inhibitors Non-nucleoside Reverse Transcriptase Inhibitors doravirine (DOR) abacavir (ABC) High-Level Resistance Potential Low-Level Resistance zidovudine (AZT) High-Level Resistance efavirenz (EFV) High-Level Resistance stavudine (D4T) High-Level Resistance etravirine (ETR) Low-Level Resistance didanosine (DDI) High-Level Resistance High-Level Resistance nevirapine (NVP) emtricitabine (FTC) High-Level Resistance rilpivirine (RPV) Low-Level Resistance lamivudine (3TC) High-Level Resistance tenofovir (TDF) Intermediate Resistance

#### RT comments

# NRTI

- D67N is a non-polymorphic TAM associated with low-level resistance to AZT.
- K70R is a TAM that confers intermediate resistance to AZT and contributes to reduced ABC and TDF susceptibility in combination with other TAMs.
- 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.
- T215Y/F are TAMs that causes intermediate/high-level resistance to AZT and potentially low-level resistance to ABC and TDF.
- . K219E/Q/N/R are accessory TAMS that usually occur in combination with multiple other TAMs.

## NNRTI

- . K101H is a non-polymorphic accessory mutation selected by NVP, EFV and ETR. When present with other NNRTI-resistance mutations, it contributes reduces susceptibility to these NNRTIs.
- V108I is a relatively non-polymorphic accessory mutation selected in vitro and/or in vivo with each of the NNRTIs. It appears to contribute to reduced susceptibility to most NNRTIs only in combination with other NNRTI-resistance mutations.
- 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

- T69N/S/A/L/E are relatively non-polymorphic mutations weakly selected in persons receiving NRTIs. They may minimally contribute reduced AZT susceptibility.
- V118I is a polymorphic accessory NRTI-resistance mutation that often occurs in combination with multiple TAMs.
- . 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.
- This virus is predicted to have low-level reduced susceptibility to RPV. The use of the combination of CAB/RPV should be considered to be relatively contraindicated.

Mutation scoring; RT

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Rule	ABC ≑	AZT ≑	D4T ≑	DDI 💠	FTC ≑	3TC ≑	TDF 💠
D67DN	5	15	15	5	0	0	5
D67DN + K70R + M184V + K219E	10	0	0	0	0	0	0
D67DN + K70R + K219E	10	15	10	10	10	10	10
D67DN + T215F + K219E	5	5	5	5	0	0	5
K70R	5	30	15	10	0	0	5
M184V	15	-10	-10	10	60	60	-10
T215F	10	60	40	15	0	0	10
K219E	5	10	10	5	0	0	5
K70R + T215F	0	0	5	5	0	0	0
Total	65	125	90	65	70	70	30

rug resista	nce mutation	Download CSV			
Rule	DOR ÷	EFV ÷	ETR ≑	NVP ≑	RPV
V108VI	10	10	0	15	0
K101H	0	10	10	15	10
G190A	0	45	10	60	15
Total	10	65	20	90	25

Drug resistance interpretation: IN

INSTI Major Mutations: E138K : G140A . S147G . Q148K

INSTI Accessory Mutations: No

IN Other Mutations: K14R - \$175N - \$125V - \$12

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#### Integrase Strand Transfer Inhibitors

bictegravir (BIC) High-Level Resistance
cabotegravir (CAB) High-Level Resistance
dolutegravir (DTG) High-Level Resistance
elvitegravir (EVG) High-Level Resistance
raltegravir (RAL) High-Level Resistance

#### IN comments

## Major

- E138K/A/T are common nonpolymorphic accessory resistance mutations selected in patients receiving RAL, EVG, CAB, and DTG. Alone they do not reduce INSTI susceptibility. However, they contribute to reduced susceptibility in combination with other mutations particularly those at position 148.
- G1405/A/C are non-polymorphic mutations that usually occur with Q148 mutations. Alone, they have minimal effects on INSTI susceptibility. However, in combination with Q148 mutations they are associated with high-level resistance to RAL and EVG and intermediate reductions in DTG and BIC susceptibility.
- \$1476 is a nonpolymorphic mutation selected in patients receiving RAL, EVG, and DTG. Alone it reduces EVG susceptibility about 5-fold.
- Q148H/K/R are nonpolymorphic mutations reported in persons receiving RAL, EVG, CAB, and DTG. They nearly always occur in combination with G140A/S or E138K. In this setting they are associated with near complete resistance to RAL and EVG, high-levels of reduction in CAB susceptibility, and low-to-intermediate reductions in DTG and BIC susceptibility.

#### Other

- 5230N is a polymorphism that is not associated with reduced INSTI susceptibility.
- There is evidence for high-level DTG resistance. If DTG is used, it should be administered twice daily.

Drug resistance mutation scores of INSTI:

Mutation scoring: IN

CSV 🖵

Rule	BIC ≑	CAB ÷	DTG ÷	EVG ‡	RAL
E138K	10	10	10	15	15
E138K + G140A	10	15	10	15	15
E138K + Q148K	10	20	10	0	0
G140A	10	10	10	30	30
G140A + Q148K	10	20	10	0	0
S147G	10	10	10	60	10
S147G+Q148K	15	20	15	0	0
<u>Q148K</u>	30	50	30	60	60
Total	105	155	105	180	130

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