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

Nucleoside Reverse Transcriptase Inhibitors

abacavir (ABC)

zidovudine (AZT)

stavudine (D4T)

didanosine (D01)

emtricitabine (FTC)

lamivudine (3TC)

tenofovir (TDF)

Susceptible

Susceptible

Susceptible

Susceptible

Susceptible

Susceptible

Non-nucleoside Reverse Transcriptase Inhibitors

doravirine (DOR)

efavirenz (EFV)

etravirine (ETR)

nevirapine (NVP)

rilpivirine (RPV)

Susceptible

Susceptible

Susceptible

Susceptible

Mutation scoring: RT

No drug resistance mutations were found for NRTL

No drug resistance mutations were found for NNRTI.

# Drug resistance interpretation: IN

HIVDB 9.5.1 (2023-11-05)

HIVDB 9.5.1 (2023-11-05)

INSTI Major Mutations: T66AI .... G118R .... E138K ....

INSTI Accessory Mutations: \$153A ===

IN Other Mutations: E11D . K14R . V31I . L45Q . V53F . E96D . L23H . V126F . V121F . V125A . V126F . V

#### 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

- . T66A/I are non-polymorphic mutations selected in persons receiving EVG, RAL, and DTG usually in combination with other INSTI-resistance mutations. They cause moderate reductions in EVG susceptibility but do not appear to reduce susceptibility to other INSTIs.
- G118R is a nonpolymorphic mutation reported in a significant proportion of persons with VF and emergent HIVDR in persons receiving a DTG-containing regimen. It has occasionally been reported with 5-10-fold reduced susceptibility to RAL, EVG, DTG and CAB, and 2-3 fold reduced susceptibility to BIC.
- 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.

## Accessory

S153Y/F are very rare mutations selected in vitro by EVG, DTG, BIC, and CAB. Alone they reduce EVG susceptibility.
 S153A is a rare mutation that alone does not appear to reduce INSTI susceptibility.

### Other

- M50I is a highly polymorphic mutation, which has a prevalence of 3% to 34% in INSTI-naïve persons depending on subtype. It has been selected in vitro by DTG and BIC in combination with R263K. It may contribute to reduced DTG and CAB susceptibility in combination with R263K.
- L74I is a highly polymorphic mutation with a prevalence of 3% to 30% depending on subtype. It is the consensus amino acid in subtype A viruses belonging to the A6 clade. It does not appear to be selected by any of the INSTIs or to reduce their susceptibility.
- V151I is an accessory INSTI selected mutation that occurs in 1% to 3% of viruses from ART-naive persons depending on subtype. Alone, it appears to have little or no effect on INSTI susceptibility.
- There is evidence for high-level DTG resistance. If DTG is used, it should be administered twice daily.

Mutation scoring: IN
HVDB 9.5.1 (2023-11-05)

Drug resistance mutation scores of INSTI:

| aray residence motorion scores or more. |       |       |       | Download Cav |       |
|---|-------|-------|-------|--------------|-------|
| Rule                                    | BIC ÷ | CAB ≑ | DTG ÷ | EVG ÷        | RAL ≑ |
| T66AI                                   | 5     | 10    | 5     | 60           | 15    |
| L74I+G118R                              | 10    | 10    | 10    | 10           | 10    |
| G118R                                   | 30    | 60    | 50    | 60           | 60    |
| G118R + E138K                           | 10    | 10    | 10    | 10           | 10    |
| E138K                                   | 10    | 10    | 10    | 15           | 15    |
| Total                                   | 65    | 100   | 85    | 155          | 110   |