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

PI Major Mutations: None

PI Accessory Mutations: None

PR Other Mutations: V11X • T12V • I13S • K14E • I15L • E35D • M36I • L38\* • R41K • H69K • K70R • L89M

#### Protease Inhibitors

Susceptible atazanavir/r (ATV/r) 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 tipranavir/r (TPV/r) Susceptible

Mutation scoring: PR

No drug resistance mutations were found for PI.

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

HIVDB 9.5.1 (2023-11-05)

NRTI Mutations: M41L • K65R • S68G • M184V

NNRTI Mutations: L1001 - K103N

RT Other Mutations: E6K • V8I • E28A • V35T • T39A • K43S • K49R • V60I • T107S • V111I • K122E • I135T • K173T • Q174K • D177E • I178L • V179I • T200A • Q207D • H208Y • R211K • K220S • Q222S • K223E • E233D • A234 • H235S • P236S • D237G • K238Q • V245E • P247Q • E248K • K249E • D250S • S251\* • W252L •

T253S • V254\* • N255Y • D256T • I257E • Q258\* • K259\* • L260K • V261I • G262N • A267G • G273X • C280V • K281\* • R284S

### **Nucleoside Reverse Transcriptase Inhibitors**

# S Non-nucleoside Reverse Transcriptase Inhibitors

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

## RT comments

# NRTI

- M41L is a TAM that usually occurs with T215Y. In combination, M41L plus T215Y confer intermediate / high-level resistance to AZT and d4T and contribute to reduced ddl, ABC and TDF susceptibility.
- K65R confers intermediate reductions in susceptibility to TDF, ABC, and 3TC/FTC. It increases AZT susceptibility. In NRTI-experienced, INSTI-naive patients with K65R, TDF+3TC+DTG. However, in 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 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.

#### 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.
- L234l 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.
- 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.
- K238T/N are uncommon non-polymorphic mutations selected in persons receiving NVP and EFV usually in combination with K103N. Alone, K238T/N appear to have minimal effects on NNRTI susceptibility. K238Q is a highly unusual mutation at this position.

Drug resistance mutation scores of NRTI:

Rule	ABC ÷	AZT \$	D4T ÷	DDI 🗦	FTC 0	3TC ≑	TDF ÷
M41L	5	15	15	10	0	0	5
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	65	-5	65	80	90	90	50

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Drug resistance mutation scores of NNK11:				Download CSV		
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	
Total	30	120	30	120	60	