PI Major Mutations: None
PI Accessory Mutations: None

PR Other Mutations: V110 • T12C • I13\* • K14\* • I15N • G16R • Q18T • E35D • M36I • R41K • K45R • R57K • L63T • E65K • H69K • L89M

# Protease Inhibitors

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

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

No drug resistance mutations were found for Pl.

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

NRTI Mutations: K65R • V75M • M184V • K219N

NNRTI Mutations: K103N • V106I • V179T • G190A

RT Other Mutations: E6N • V8I • V35T • V60I • S68K • K122E • D123N • I135T • K173L • Q174K • D177E • T200A • I202V • Q207A • R211S • K238X • V245Q • P247X • Δ263-264 • W266\* • A267M • S268G • Q269S • I270Q • Y271F • A272M • G273Q • I274D • Q278H • L279W • K281X • T286A • L289P • T290N • E291R • V292R • I293W

# Nucleoside Reverse Transcriptase Inhibitors

# Non-nucleoside Reverse Transcriptase Inhibitors

abacavir (ABC)	High-Level Resistance	doravirine (DOR)	Potential Low-Level Resistance
zidovudine (AZT)	Susceptible	efavirenz (EFV)	High-Level Resistance
stavudine (D4T)	High-Level Resistance	etravirine (ETR)	Low-Level Resistance
didanosine (DDI)	High-Level Resistance	nevirapine (NVP)	High-Level Resistance
emtricitabine (FTC)	High-Level Resistance	rilpivirine (RPV)	Low-Level Resistance
lamivudine (3TC)	High-Level Resistance		
tenofovir (TDF)	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.
- V75T/M/A/S are nonpolymorphic accessory NRTI-selected mutations. They appear to have minimal phenotypic effects on AZT, ABC, and TDF.
- M184V/I cause high-level in vitro resistance to ATC 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.
- K219E/Q/N/R are accessory TAMS that usually occur in combination with multiple other TAMs.

# NNRTI

- K103N is a non-polymorphic mutation that confers high-level reductions in NVP and EFV susceptibility. It is the most commonly transmitted DRM.
- V106I occurs in 1% to 2% of viruses from untreated persons. It contributes to reduced NNRTI susceptibility only in combination with other NNRTI-resistance mutations. It is commonly selected in persons receiving DOR in combination with mutations at position 227.
- V179T is a rare non-polymorphic mutation occasionally selected in persons receiving NNRTIs. It is associated with minimal, if any, reduction in ETR and RPV susceptibility.
- 6190A 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.
- 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.

Drug	resist	ance	mutation	scores	ĺ

Download CSV s of NRTI: ABC 
AZT D4T 
DDI 
TTC 
TTC 
TTC K65R 30 60 60

90

90

Drug resistance mutation scores of NNRTI:

90

45

Rule	DOR \$	EFV \$ ETR \$		NVP \$	RPV	
<u>V106I</u>	10	0	10	10	10	
K103N	0	60	0	60	0	
G190A	0	45	10	60	15	
Total	10	105	20	130	25	

PI Major Mutations: None
PI Accessory Mutations: None

PR Other Mutations: T12M • I13\* • K14\* • E21X • E35D • M36I • R41K • R57K • L63P • H69K • K70R • L89M

## Protease Inhibitors

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

Mutation scoring: PR

No drug resistance mutations were found for Pl.

HIVDB 9.5.1 (2023-11-05)

HIVDB 9.5.1 (2023-11-05)

Drug resistance interpretation: RT

NRTI Mutations: M184V

NNRTI Mutations: K101H • G190A • F227I

RT Other Mutations: K11T - K20R - V21I - V35T - T39R - K122E - D123N - I135T - I142T - K173S - D177G - V179I - Q207N - R211X - Q222H - K223Q - E224K - P225N - L228S - P236S - D237\* - K238Q - V245Q - P247Q - E248T - K249R - D250E - S251L - W252T - T253V - V254M - N255I - D256Y - I257R

# Nucleoside Reverse Transcriptase Inhibitors

abacavir (ABC) Low-Level Resistance
zidovudine (AZT) Susceptible
stavudine (D4T) Susceptible

didanosine (DDI) Potential Low-Level Resistance
emtricitabine (FTC) High-Level Resistance
lamivudine (3TC) High-Level Resistance

tenofovir (TDF) Susceptible

# Non-nucleoside Reverse Transcriptase Inhibitors

doravirine (DOR) High-Level Resistance
efavirenz (EFV) High-Level Resistance
etravirine (ETR) Low-Level Resistance
nevirapine (NVP) High-Level Resistance
rilpivirine (RPV) Low-Level Resistance

# RT comments

# NRTI

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.

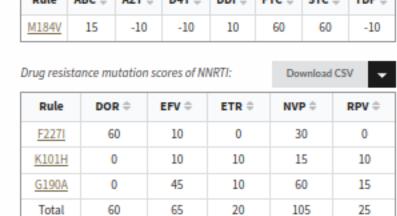
# NNRTI

- 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.
- 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.
- F227L is a non-polymorphic mutation that usually occurs in combination with V106A. It is selected in vivo and in vitro with both NVP and DOR. In this context it is associated with high-level reductions in NVP and DOR.
   F227L is a non-polymorphic mutation that usually occurs in combination with V106A. It is selected in vitro by DOR.

# 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.
- P225H is a non-polymorphic EFV-selected mutation that usually occurs in combination with K103N. The combination of P225H and K103N synergistically reduces NVP, EFV and DOR susceptibility. P225N 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.
- 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.
- 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.

HIVDB 9.5.1 (2023-11-05)



RPV

15

25

PI Major Mutations: None PI Accessory Mutations: None

PR Other Mutations: L10G • V11\* • T12V • I13S • K14N • I15D • G16R • Q18M • K20Q • L33V • M36I • R41K • L63P • I64V • I72V

# Protease Inhibitors

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

# PR comments

# Other

- L10F is a common non-polymorphic, PI-selected accessory mutation associated with reduced in vitro susceptibility to LPV and DRV. L10I/V are polymorphic, PI-selected accessory mutations. Their effects on PI susceptibility have not been well studied. L10G is a highly unusual mutation at this position.
- . L33I/V are minimally polymorphic mutations that do not appear to be selected by PIs or to reduce their susceptibility.

# Mutation scoring: PR

No drug resistance mutations were found for Pl.

Drug resistance interpretation: RT

HIVDB 9.5.1 (2023-11-05)

HIVDB 9.5.1 (2023-11-05)

NRTI Mutations: K70Q • M184I

NNRTI Mutations: K101E • K103N • G190A

RT Other Mutations: V35I • T39E • V60I • V90I • S105T • D121Y • K122E • I135K • Q174R • D177E • I178V • T200I • Q207E • R211K • K219X • P225X • P226S • L228R • P236S • L246T • P247A • E248R • D250E • N255M • D256I • I257Y • Q258R • K259V • L260V • V261E • G262N • L264W

# **Nucleoside Reverse Transcriptase Inhibitors**

# abacavir (ABC) zidovudine (AZT) stavudine (D4T) didanosine (DDI) emtricitabine (FTC) lamivudine (3TC) tenofovir (TDF) Intermediate Resistance Low-Level Resistance Low-Level Resistance High-Level Resistance Low-Level Resistance

# Non-nucleoside Reverse Transcriptase Inhibitors

doravirine (DOR)
Low-Level Resistance
efavirenz (EFV)
High-Level Resistance
etravirine (ETR)
Intermediate Resistance
nevirapine (NVP)
High-Level Resistance
rilpivirine (RPV)
High-Level Resistance

# RT comments

# NRTI

- 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

- K101E is a non-polymorphic accessory mutation that confers intermediate resistance to NVP and RPV and low-level reductions in susceptibility to EFV, ETR, and DOR when it occurs with other NNRTI-resistance mutations.
- K103N is a non-polymorphic mutation that confers high-level reductions in NVP and EFV susceptibility. It is the most commonly transmitted DRM.
- 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

- 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.
- . 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.

Drug	resistance	mutation	scores of	NRTI:



Rule	ABC \$	AZT \$	D4T \$	DDI \$	FTC \$	зтс ≑	TDF
<u>K700</u>	15	0	15	15	10	10	15
M184I	15	-10	-10	10	60	60	-10
K70Q + M184I	0	0	10	0	0	0	10
Total	30	-10	15	25	70	70	15





Drug resistance mut	ation score	s of NNRTI:		Download C	SV
Rule	DOR \$	EFV \$	ETR \$	NVP \$	RPV
<u>K101E</u>	15	15	15	30	45
K101E + G190A	5	0	5	0	0
<u>K103N</u>	0	60	0	60	0
<u>G190A</u>	0	45	10	60	15
K101E + M184I	0	0	0	0	15
Total	20	120	30	150	75

PI Major Mutations: None
PI Accessory Mutations: K20T

PR Other Mutations: L10G • V11N • T12F • I13S • G16C • G17R • Q18K • E35D • M36I • R41K • K45R • R57K • H69K • L89M

# Protease Inhibitors

atazanavir/r (ATV/r)
darunavir/r (DRV/r)
fosamprenavir/r (FPV/r)
indinavir/r (IDV/r)
lopinavir/r (LPV/r)
Susceptible
Susceptible
Susceptible
Susceptible

nelfinavir (NFV) Low-Level Resistance

saquinavir/r (SQV/r) Susceptible tipranavir/r (TPV/r) Susceptible

#### PR comments

## Accessory

K20T is a non-polymorphic accessory PI-selected mutation associated with reduced susceptibility to ATV and LPV.

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# Other

• L10F is a common non-polymorphic, PI-selected accessory mutation associated with reduced in vitro susceptibility to LPV and DRV. L10I/V are polymorphic, PI-selected accessory mutations. Their effects on PI susceptibility have not been well studied. **L10G** is a highly unusual mutation at this position.

Drug resistance mutation scores of PI:

Mutation scoring: PR

HIVDB 9.5.1 (2023-11-05)

Ding it								· ·
Rule	ATV/r 🕏	DRV/r 🕏	FPV/r 🗢	IDV/r 🗢	LPV/r ≑	NFV \$	sqv/r =	TPV/r 🗢
K20T	5	0	5	5	0	15	5	0

# Drug resistance interpretation: RT

HIVDB 9.5.1 (2023-11-05)

NRTI Mutations: M184V NNRTI Mutations: K103N

RT Other Mutations: P4Q · K11T · K20R · V21I · E28K · K32E · V35T · T39N · V60I · K101A · K122E · D123N · I135T · K173A · Q174K · I195X · T200A · Q207E · R211K · K219X · P236S · D237\* · K238Q · V245Q · P247Q · D250E · S251C · T253N · V254C · N255H · L260S · V261S · L264I · N265X · A267V · A272P · I274S ·

K275S - V276K - K277A - Q278L - L279C - C280R - K281T - L283S - R284D - G285R - T286S - K287L - A288K - L289P - T290L - E291N - V292R - I293M

# **Nucleoside Reverse Transcriptase Inhibitors**

# Non-nucleoside Reverse Transcriptase Inhibitors

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

tenofovir (TDF) High-Level F Susceptible

#### RT comments

## NRTI

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

K103N is a non-polymorphic mutation that confers high-level reductions in NVP and EFV susceptibility. It is the most commonly transmitted DRM.

# Other

- K101N/A/T are uncommon non-polymorphic NNRTI-selected mutation of uncertain phenotypic and clinical significance.
- 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.

# Mutation scoring: RT

Drug resistance mutation scores of NRTI:

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Rule	ABC \$	AZT \$	D4T ‡	DDI \$	FTC 0	зтс ≑	TDF (
M184V	15	-10	-10	10	60	60	-10

Drug resistance mutation scores of NNRTI:

Download CSV

Rule	DOR \$	EFV \$	ETR \$	NVP ≎	RPV =
K103N	0	60	0	60	0

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HIVDB 9.5.1 (2023-11-05)

PI Major Mutations: None PI Accessory Mutations: None

Drug resistance interpretation: PR

PR Other Mutations: V11W - T12L - I13A - K14Q - I15S - G16R - Q18K - L19I - K20H - E35D - M36I - R41K - R57K - H69K - L89M

# **Protease Inhibitors**

atazanavir/r (ATV/r) Susceptible darunavir/r (DRV/r) Susceptible Susceptible fosamprenavir/r (FPV/r) 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 Pl.

Drug resistance interpretation: RT

M41L · E44D · S68G · L74I · M184V · L210W · T215Y · K219N NRTI Mutations:

NNRTI Mutations: A98G - K103N - V108I - H221Y

RT Other Mutations: E6D • K20R • V35T • V60I • V118I • K122E • D123N • I142V • S162N • K166R • F171Y • K173A • Q174K • D177E • I178M • V179I • T200A • I222V • E203D • Q207A • R211N • K223X • P226H • L234X • P236S • K238X • T240D • V241S • P243S • V245Q • P247Q • K249Q • N255M • D256I • I257Y • Q258R • K259N •

L260S • V261G • G262K • K263L • L264M • ∆265-266 • A267X • S268Q • Q269S • A272S • K277E • Q278A • L279T • C280G • K281V • L282N • L283F • R284F • G285K

# Nucleoside Reverse Transcriptase Inhibitors

# Non-nucleoside Reverse Transcriptase Inhibitors

abacavir (ABC)	High-Level Resistance	doravirine (DOR)	Intermediate 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	nevirapine (NVP)	High-Level Resistance
emtricitabine (FTC)	High-Level Resistance	rilpivirine (RPV)	Intermediate Resistance
lamivudine (3TC)	High-Level Resistance		
tenofovir (TDF)	High-Level 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.
- E44D is a relatively non-polymorphic accessory mutation; E44A is a nonpolymorphic accessory mutation. Each usually occurs with multiple TAMs.
- \$686 is a polymorphic mutation that is often selected in combination with K65R. It partially restores the replication defect associated with K65R.
- L74V causes intermediate ABC resistance. L74I causes low-level ABC resistance.
- 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.
- . L210W is a TAM that usually occurs in combination with M41L and T215Y. The combination of M41, L210W and T215Y causes high-level resistance to AZT and intermediate resistance to ABC and TDF.
- 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

- A986 is a non-polymorphic accessory mutation associated with low-level reduced susceptibility to each of the NNRTIs.
- K103N is a non-polymorphic mutation that confers high-level reductions in NVP and EFV susceptibility. It is the most commonly transmitted DRM.
- 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.
- H221Y is a non-polymorphic accessory mutation selected primarily by NVP, RPV, and DOR. It frequently occurs in combination with Y181C.

# Other

- 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.
- 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.
- This virus is predicted to have intermediate-level reduced susceptibility to RPV. The use of the combination of CAB/RPV should be considered to be contraindicated.

Drug resistance mutation scores of NRTI:

Mutation scoring: RT

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Rule	ABC ≑	AZT ≑	D4T ≑	DDI \$	FTC ÷	зтс ≑	TDF 0
M41L	5	15	15	10	0	0	5
M41L + E44D + L210W + T215Y	5	5	5	5	0	0	5
M41L + M184V + T215Y	10	0	0	0	0	0	0
M41L + L210W	10	10	10	10	0	0	10
M41L + L210W + T215Y	10	0	0	0	15	15	10
M41L + T215Y	10	10	10	10	5	5	10
<u>L741</u>	15	0	0	60	0	0	5
M184V	15	-10	-10	10	60	60	-10
<u>L210W</u>	5	15	15	10	0	0	5
L210W + T215Y	10	10	10	10	0	0	10
<u>T215Y</u>	10	60	40	15	0	0	10
K219N	5	10	10	5	0	0	5
Total	110	125	105	145	80	80	65

HIVDB 9.5.1 (2023-11-05)

Drug resisto	ince mutatio	n scores of N	NRTI:	Download	CSV
Rule	DOR ÷	EFV \$	ETR ‡	NVP ≎	RPV ≎
<u>A98G</u>	15	15	10	30	15
V108I	10	10	0	15	0
HOOTY	10	10	10	15	15

PI Major Mutations: None
PI Accessory Mutations: None

PR Other Mutations: V11G • T12S • I13S • K14R • L33V • M36I • N37D • P39Q • R41K • R57K • D60E • I62V • L63P • I64V

# Protease Inhibitors

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

# PR comments

# Other

. L331/V are minimally polymorphic mutations that do not appear to be selected by PIs or to reduce their susceptibility.

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

No drug resistance mutations were found for PI.

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

NRTI Mutations: L74I - M184V

NNRTI Mutations: K103N • E138G • V179L • H221Y

RT Other Mutations: K20R • V35I • K49R • V60I • K102H • D121Y • K122E • I135T • S162C • D177E • I178L • T200A • R206K • Q207G • R211K • D237E • \( \triangle \triangle

# Nucleoside Reverse Transcriptase Inhibitors

zidovudine (AZT)
stavudine (D4T)
didanosine (DDI)
emtricitabine (FTC)
lamivudine (3TC)
tenofovir (TDF)

Susceptible
High-Level Resistance
High-Level Resistance
Susceptible

# Non-nucleoside Reverse Transcriptase Inhibitors

doravirine (DOR)
Potential Low-Level Resistance
efavirenz (EFV)
High-Level Resistance
Intermediate Resistance
nevirapine (NVP)
High-Level Resistance
Intermediate Resistance
Intermediate Resistance

# RT comments

abacavir (ABC)

# NRTI

L74V causes intermediate ABC resistance. L74I causes low-level ABC resistance.

Intermediate Resistance

M184V/I cause high-level in vitro resistance to ATC 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

- 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.
- V179L is a rare non-polymorphic mutation listed as a RPV-associated resistance mutation by the FDA package insert. Its effects on NNRTI susceptibility have not been well studied.
- H221Y is a non-polymorphic accessory mutation selected primarily by NVP, RPV, and DOR. It frequently occurs in combination with Y181C.
- This virus is predicted to have intermediate-level reduced susceptibility to RPV. The use of the combination of CAB/RPV should be considered to be contraindicated.

#### RT comments

# NRTI

- L74V causes intermediate ABC resistance. L74I causes low-level ABC resistance.
- 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

- . 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.
- . V179L is a rare non-polymorphic mutation listed as a RPV-associated resistance mutation by the FDA package insert. Its effects on NNRTI susceptibility have not been well studied.
- . H221Y is a non-polymorphic accessory mutation selected primarily by NVP, RPV, and DOR. It frequently occurs in combination with Y181C.
- . This virus is predicted to have intermediate-level reduced susceptibility to RPV. The use of the combination of CAB/RPV should be considered to be contraindicated.

Mutation scoring: RT

HIVDB 9.5.1 (2023-11-05)

Drug resistance mutation scores of NRTI:



Rule	ABC ≎	AZT ≎	D4T ≎	DDI 🗘	FTC ‡	зтс ≑	TDF 🗢
<u>L741</u>	15	0	0	60	0	0	5
M184V	15	-10	-10	10	60	60	-10
Total	30	-10	-10	70	60	60	-5

Drug resistance mutation scores of NNRTI:

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Rule	DOR =	EFV \$	ETR ‡	NVP ≑	RPV \$
<u>H221Y</u>	10	10	10	15	15
K103N	0	60	0	60	0
E138G	0	10	10	10	15
<u>V179L</u>	0	10	10	10	15
Total	10	90	30	95	45