

## An Approach to Treatment Failure in adults with HIV

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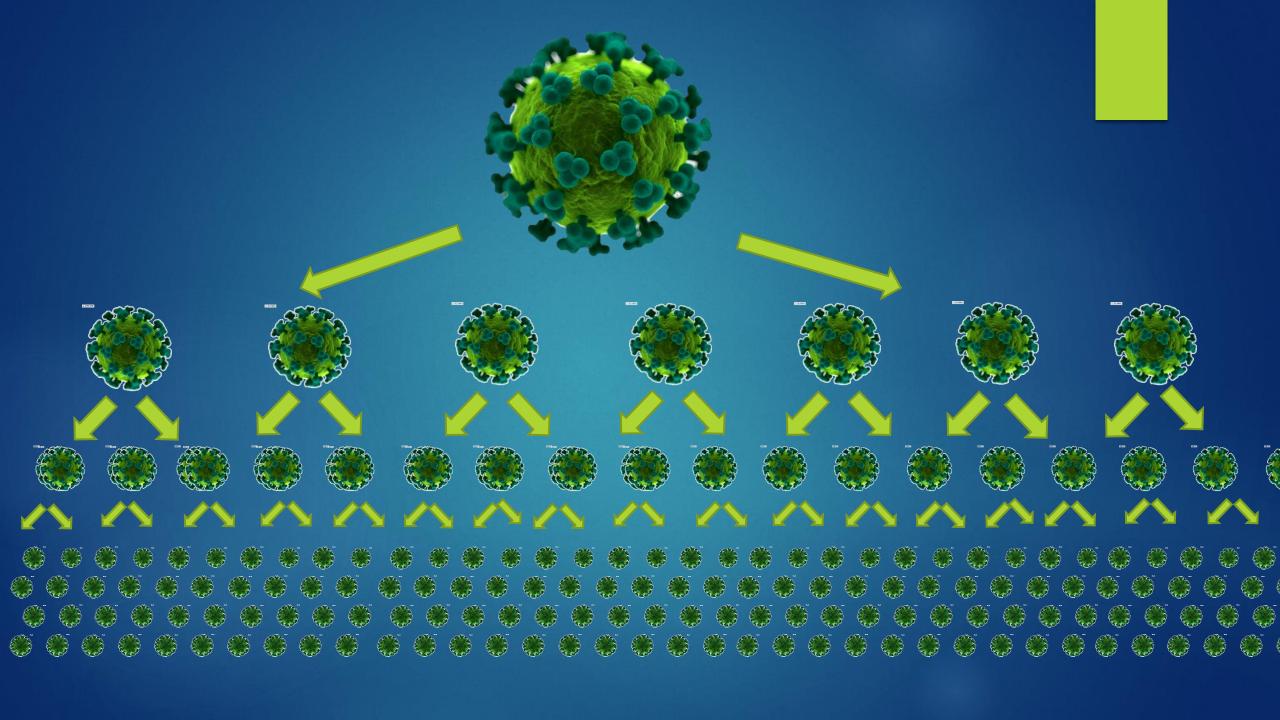
### Where does the HIV virus live?

- ▶ In the Blood?
- **×**No
- ► Lymph nodes?
- Yes
- ▶ In the liver?
- Yes
- ► Spleen?
- Yes



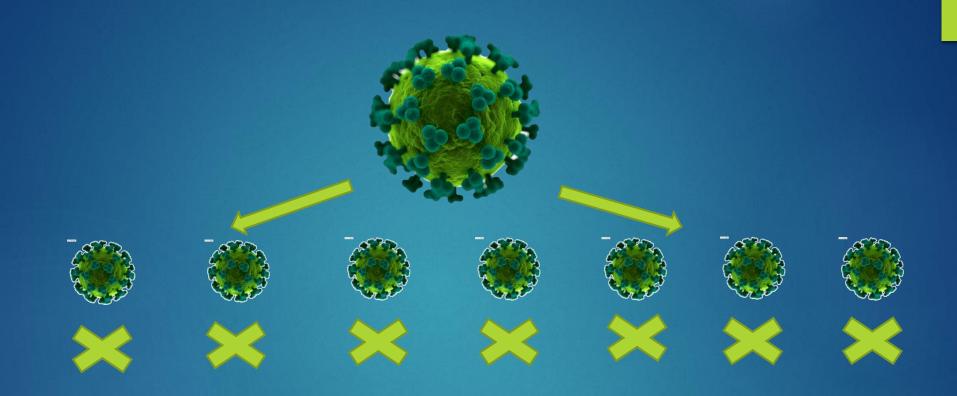
## What does a Viral Load tell us?





## What does an undetectable Viral Load tell us?



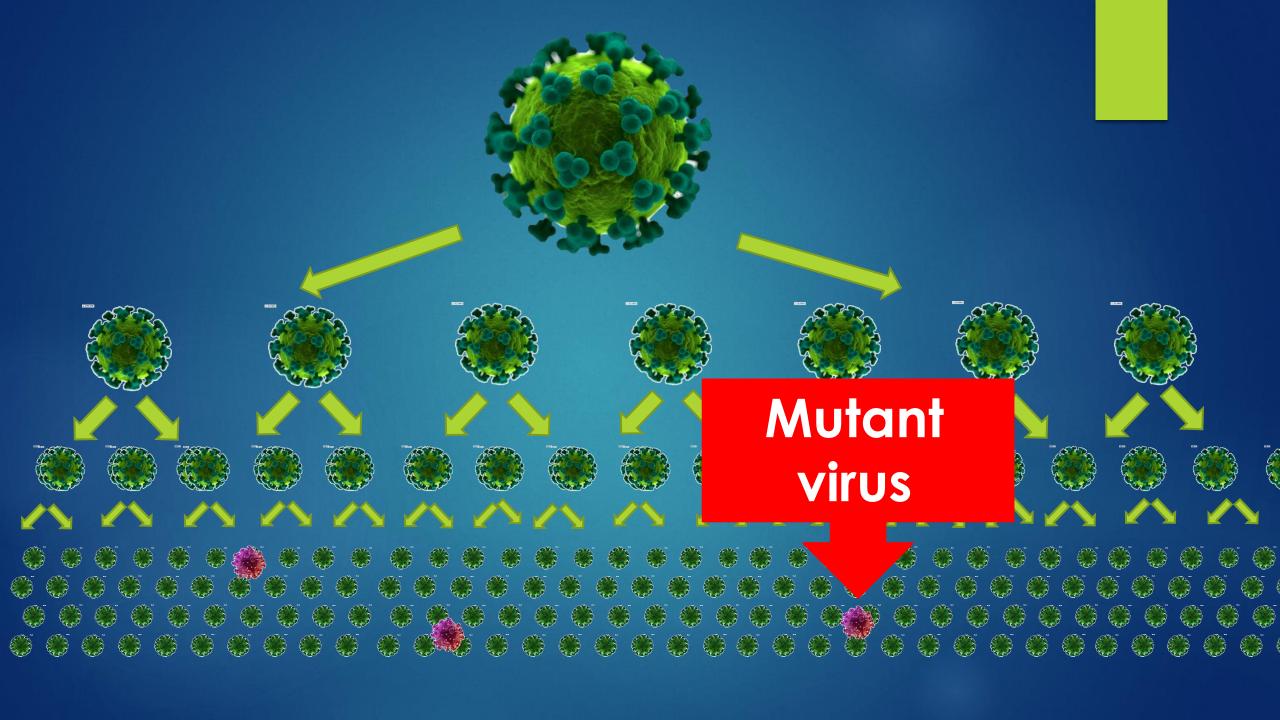


# An Undetectable Viral load means that the ARVS are working and the virus is not multiplying

- So HIV cant overflow into the blood stream
- ► HIV will not kill off CD4 cells
- ▶ The patient will not get sick
- ▶ The patient will not transmit HIV to their baby
- ▶ The patient will not transmit HIV to their sexual partner

The Partner Study

# What is ARV Resistance?

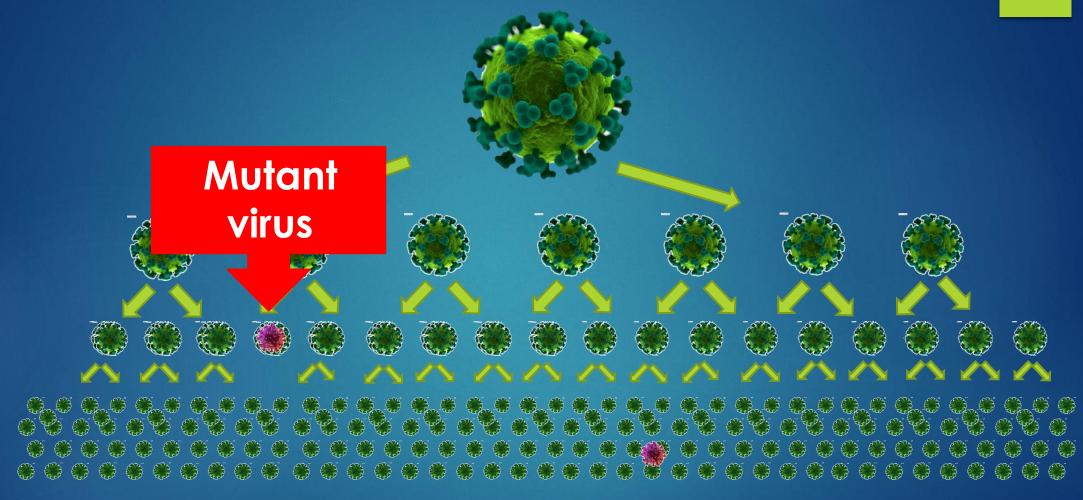


## We need 2 things for resistance to develop

1) Viral multiplication (replication)

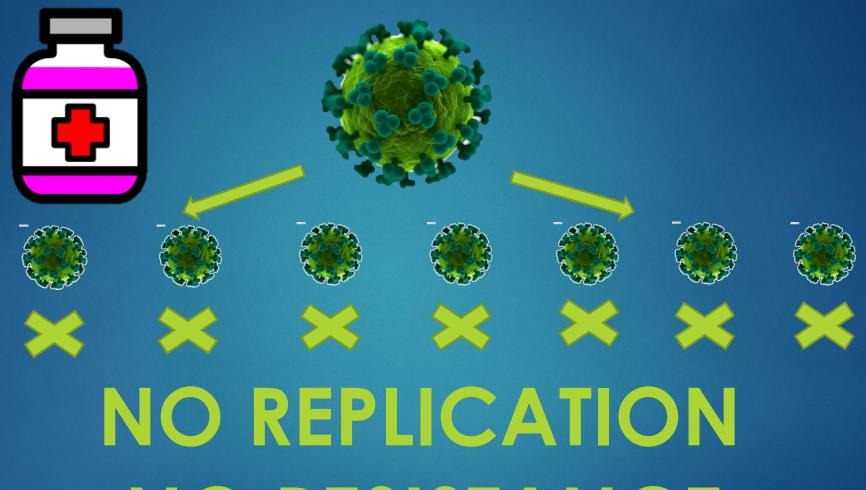
2)Low levels of ARVs in the body

### NO ARVS



## NO RESISTANCE

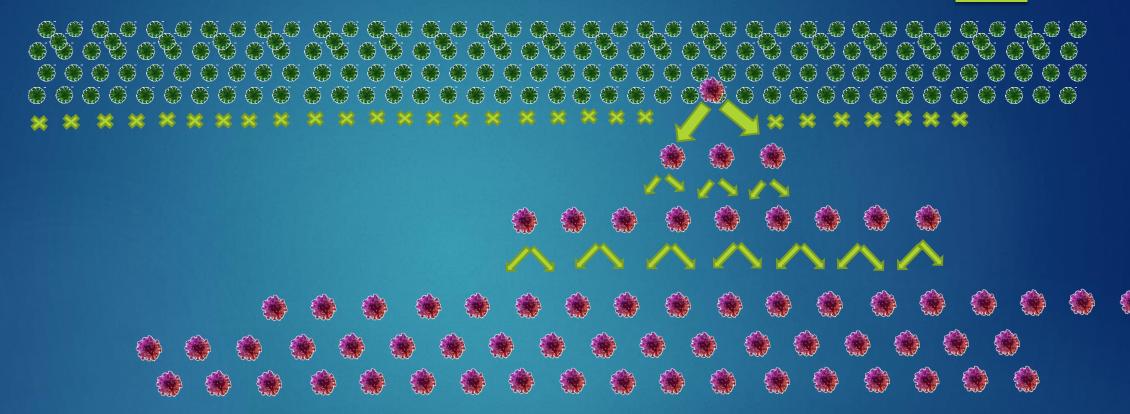
### SUFFICIENT LEVELS OF ARVS



NO RESISTANCE



### LOW LEVELS OF ARVS PRESENT



## REPLICATION + LOW LEVELS OF ARVS =RESISTANCE

# What will happen to this person?

They will need to change ARV's

We need to prevent them from failing first line!

### What can lead to resistance?

Anything which decreases the amount of ARVs in your body eg:

- Missing doses of ARVs
- Incorrect dosing
- Drug interactions which decrease the amount of ARV's in your body eg. Rifampicin with LPV/r or DTG
- Severe prolonged vomiting or diarrhoea/ malabsorption

Transmitted resistance

### Approach to high VLs:

VL is suppressed

**VL<50** 

Not replicating
Sufficient levels of
effective ARVs

Low level viraemia

VL: 50-1000

- Viral blip
- Persistant low level viraemia

**Viral Failure** 

VL>1000

- Not adherent
- Inadequate drug levels
- Resistance
- Or multiple

## What is a viral blip?

- Small Increase in viral load between 50 and < 1000 copies/ml</p>
- Caused by:
  - Normal burst of viral replication even if treatment is working
- Does not affect how ARVs work
- Does not cause resistance
- Only way to confirm that it is a blip is to repeat the viral load after 3 months and if the repeat viral load is <50 copies/ml then it was a blip

**VL>1000** 

Not adherent

Inadequate drug levels

Resistant

Assessadherence

- Correct doses?
- Drug interactions?

### VL Monitoring Algorithm for Clients on TLD

(also applicable to ALD and other DTG-containing regimens)

Routine VL monitoring for virally suppressed clients as outlined in the algorithm 
"Routine HIV VL Monitoring on ART" on page 20

VL monitoring should happen every 6 DCs in a breastfeeding woman

VL < 50 c/mL

VL unsuppressed (VL ≥ 50 c/ml)
(This includes previous VL level of 50-999 and VL > 1000 c/ml)



Do a thorough assessment of the cause of an elevated VL. Consider the possibility of:

- A. Adherence problems (see "Enhanced Adherence Support" on page 22)
- B. <u>Bugs</u> (Intercurrent infections)
- C. In-Correct ART dosage (see Annexure 5 "Drug Dosing Chart" on page 34)
- D. <u>Drug Interactions</u> (see"Drug Interactions with DTG and Rifampicin-containing TB Treatment" on page 13)
- E. REsistance (if > 2 years on treatment)

Implement interventions to re-suppress the VL, including Enhanced Adherence Support if indicated (See Annexure 3 Enhanced Adherence Counselling)

Recommend condom use and contraception as appropriate

Repeat VL after 3 months

# If VL>50 do ABCDE

### Assessing an Elevated Viral Load A thorough assessment is essential for any client with a viral load measuring ≥ 50 c/ml Is adherence to medication poor? Ask about factors that may influence adherence e.g. Direct cost of clinic visits to patient, e.g. transport, loss of income, cost of paying another Tips person to take on social responsibilities Ask open ended questions e.g. . Taking time away from existing work, finding work and/or social care responsibilities 'What makes it difficult for you to · Needing to travel for extended periods of time collect or take your treatment?", Medication side-effects and "How many doses have you missed this week?" **A** Adherence Statements like "we all miss a dose now and then" can encourage a client to be more open. Pregnant women may experience nausea/vomitting, heartburn, and constipation. Assess the need for symptomatic treatment with an anti-emetic, anti-diarrhea agent, or fiber Create a safe and nonsupplement. judgemental space for your client Adherence difficulties in young children are often linked to poor tolerability of to discuss challenges. unpalatable formulations, particularly LPV/r solution. It is important to ask the caregiver about how the child tolerates the medication e.g., does the child refuse to swallow the Remember that immune Bugs compromised, malnourished, and pregnant clients may not exhibit overt symptoms of TB. If in doubt, do a TB GXP. Correct dose I function or previous renal impairment See also "Drug Interactions with DTG and Rifampicin-containing TB Treatment" on page 13 **Drug interactions** If in any doubt, call the HIV Hotline 0800 212 506 or one of the "Helplines" on page 23 Refer to the algorithm rEsistance "Management of Confirmed ded and Virological Failure on TLD" on page 23

How do you assess adherence?

Do patients tell you the truth?



I'm unemployed and drink alcohol with friends to pass the time, I usually drink so much I don't remember anything for 3-5 days of the week



### Enhanced Adherence Counselling

### Clinician considerations for providing Enhanced Adherence Counselling (EAC)

| Barrier to adherence                                     | Intervention   | EAC indicated?  |
|--|--|-----------------|
| Difficulty getting to facility to collect treatment      | Reduce unnecessary visits through enrolling client in a RPCs model or providing multi-month dispensing (MMD) | No need for EAC |
| Drug side effects or unpalatability impacting adherence? | Change to more palatable regimen   | No need for EAC |
| Challenges with taking/remembering to take treatment     | Provide EAC  |                 |

### Enhanced Adherence Support

Enhanced Adherence Counselling (EAC) is aimed at non-stable clients presenting with adherence issues or poor treatment response and/or signs of treatment failure. Enhance Adherence Counselling focuses on:

- Providing education on the outcome of their latest clinical assessment and VL results
- Understanding what the client already knows or doesn't know regarding their treatment and the importance of VL suppression
- · Doing a mental health screen
- Correcting any misconceptions and allowing flexibility around the most common barriers to adherence (such as alcohol/ drug consumption, forgetting doses due to a rigid schedule, etc.).
- · Assessing and understanding the barriers that affect the client's adherence
- Developing adherence strategies to overcome these

To support the above processes, the following useful tools extracted from the Differentiated Care Models Standard Operating Procedures 2023 included in the annexures:

- SOP 2 Enhanced Adherence Counselling (Annexure 3)
- Mental Health Screen (Annexure 4)
- . Child and adolescent disclosure counseling for children living with HIV (Annexure 7)

'better late than never': clients should be counselled they can take their ARVs up to several hours late if they miss their chosen time



### More clues

If you see 2 patients with the following viral loads, which one do you think is more likely to have resistance?

▶ VL: 2 500

VL: 800 000

Resistance mutations generally weaken the HIV virus and reduce its replicability, but it is not always the case.



## Some ARVs are easy to develop resistance to and some are difficult



Quickly develops resistance

3TC / NVP / EFV

Takes a long time to develop resistance (> 2 years)

LPV/r & DTG

# Approach to high VL on EFV/NVP

### Switch to TLD

### NADIA trial

- In Patients failing TDF/3TC/NNRTI regimen TDF/3TC/DTG (TLD )was superior to AZT/3TC/DTG in 2nd line
- ► ARTIST, VISEND and D<sup>2</sup>EFT Trials show similar results

### New ART Guidelines:

## Switch all patients on EFV/NVP to TLD regardless of VL

| VL<br>considerations                    | Current Regimen  | Criteria for switch   | Regimen if change indicated  |
|---|--|---|--|
| Switching<br>regardless of VL<br>result | TEE  ABC/3TC/EFV (or NVP*)  AZT/3TC/EFV (or NVP*)  AZT/3TC/DTG  Any LPV/r or ATV/r regimen for less than 2 years | Switch all to a DTG-containing regimen, regardless of VL result  Review VL in last 12 months.  If VL in last 12 months was not suppressed, continue to switch same day, but do ABCDE assessment and provide enhanced adherence counseling (EAC) if needed.  If VL was not done in last 12 months, do it at this visit, but do not wait for the result to switch | TLD  provided no renal dysfunction and age ≥ 10 yrs and weight ≥ 30 kg  If client does not qualify for TDF  ABC¹/3TC/DTG  If client does not qualify for TDF and has ABC hypersensitivity  AZT/3TC/DTG |

ABC data Vs TDF data

TLD 1

Clients on a DTG-containing regimen, who have never failed a previous regimen (old "1st line" terminology)

TLD 2

Clients on a DTG-containing regimen, who have failed a previous regimen (old "2nd line" terminology)

Approach to high VL on a Pl (LPV/r or ATV/r) HIV

Takes a long time to develop resistance (> 2 years)

LPV/r & DTG

### New Guidelines: If on PI < 2 years: switch to TLD

### (Non VL-dependent regimen switches)

| VL<br>considerations                    | Current Regimen   | Criteria for switch   | Regimen if change indicated  |
|---|---|---|--|
| Switching<br>regardless of VL<br>result | ABC/3TC/EFV (or NVP*)  AZT/3TC/EFV (or NVP*)  AZT/3TC/DTG  Any LPV/r or ATV/r regimen for less than 2 years PTO | Switch all to a DTG-containing regimen, regardless of VL result  Review VL in last 12 months.  If VL in last 12 months was not suppressed, continue to switch same day, but do ABCDE assessment and provide enhanced adherence counseling (EAC) if needed.  If VL was not done in last 12 months, do it at this visit, but do not wait for the result to switch | TLD  provided no renai dysfunction and age ≥ 10 yrs and weight ≥ 30 kg  If client does not qualify for TDF  ABC¹/3TC/DTG  If client does not qualify for TDF and has ABC hypersensitivity  AZT/3TC/DTG |

ABC data Vs TDF data

## If on a PI > 2 years: VL dependent switches

- Look at their VL result in the last 12 months
- If they are failing a PI regimen, they might have PI resistance which means that they have no "backup" regimen if they fail DTG
- They may require a resistance test to determine if they indeed have PI resistance
- If resistance is confirmed, they will require an individualised regimen (to be determined in consultation with an expert)

Switching Existing Clients to DTG-containing Regimens
(Adults, adolescents or children who have never used a DTG-containing regimen in the past)

| <b>VL-dependent regimen switches</b><br>Relevant to all clients who have been on PI-based regimens for more than two years: their VL result in the last 12 months<br>will influence the decision of how and when to switch to a DTG-containing regimen |  |  |   |  |
|--|--|--|---|--|
| VL<br>considerations   | Current Regimen  | Criteria for switch  | Regimen if change indicated   |  |
| VL < 1000 c/mL   | Any<br>LPV/r or ATV/r<br>regimen for<br>more than 2 years                                  | Switch all to a DTG-containing regimen If VL in last 12 months was ≥ 50 c/mL, continue to switch same day, but do ABCDE assessment, provide EAC if needed, and repeat the VL after 3 months as per "The VL non-suppression algorithm" on page 19                         | TLD  provided no renal dysfunct on and age ≥ 10 yrs and weight ≥ 30 kg  If clients does not qualify for TDF  ABC¹/3TC/DTG |  |
|  | Adult or adolescent on any LPV/s or ATV/s regimen and adherence less than 80% <sup>3</sup> | Do not do a resistance test  These clients are unlikely to have PI resistance mutations. Rather switch to a more tolerable once using FDC regimen which is likely to support adherence. Manage as per "The VL non-suppression algorithm" on page 19                      | TLD provided in a remaining for and age ≥ 10 yrs and weight ≥ 30 kg  If clients does not qualify for TDF ABC¹/3TC/DTG     |  |
| <sup>2</sup> Two or more VLs ≥ 1000 c/mL taken two or more years after starting PI regimen   | Adult or adolescent on any regimen and adherence more than 80% <sup>3</sup>                | Clients who meet the definition of co<br>and have confirmed adherence more than<br>Discuss with an HIV expert <sup>4</sup> to authorise<br>Provide individualised regimen as rec<br>Repeat VL 3 months after the regimen chang<br>the "Management of Confirmed Virologic | and interpret a resistance test.  commended by HIV expert. e to confirm re-suppression, as per                            |  |
|  | Child < 10 years,<br>or weight < 30 kg<br>on any<br>LPV/r or ATV/r regimen                 | These clients do not yet qualify for TLD and may require a resistance test.  Refer to algorithm "Switching children on PI-containing regimens to DTG-containing regimens" on page 16   |   |  |

# Fine print:

- Objective measures of good adherence include at least one of:
  - ▶ Pharmacy refills > 80% in the last 6-12 months (if this is known)
  - Attendance of > 80% of scheduled clinic visits in the last 6-12 months (if this is known)
  - Detection of current antiretroviral drug/s in the client's blood or urine, if available
  - Note: Self-reported adherence is not considered a reliable measure of good adherence

# If there is PI resistance

### Third Line ART:

- ► Likely TLD if DRV is fully susceptible
- Discuss with an expert

# Approach to high VL on DTG

### **VL Monitoring for Clients on TLD**

(also applicable to ALD and other DTG-containing regimens)

Routine VL monitoring for virally suppressed clients as outlined in the algorithm "Routine HIV VL Monitoring on ART" on page 18



VL < 50 c/mL

VL unsuppressed (VL ≥ 50 c/ml) (This includes previous VL level of 50-999 and VL > 1000 c/ml)



Do a thorough assessment of the cause of an elevated VL. Consider the possibility of:

- A. Adherence problems (see "Enhanced Adherence Support" on page 20)
- B. **B**ugs (Intercurrent infections)
- C. In-Correct ART dosage (see Annexure 5 "Drug Dosing Chart" on page 34)
- D. <u>Drug Interactions</u> (see "Drug Interactions with DTG and Rifampicin-containing TB Treatment" on page 13)
- E. REsistance (if > 2 years on treatment)

Implement interventions to re-suppress the VL, including Enhanced Adherence Support if indicated (See Annexure 3 Enhanced Adherence Counselling)

Recommend condom use and contraception as appropriate

Repeat VL after 3 months

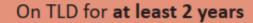
### Repeat VL after 3 months

### Repeat VL unsuppressed 1 (VL > 50 c/ml)



### Re-assess and resolve adherence issues! 2

(See "ABCDE assessment of an Elevated Viral Load" on page 20 and Annexure 3 "Enhanced Adherence Support" on page 20)



On TLD less than 2 years <sup>3</sup>

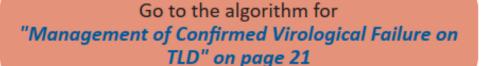


If Adherence > 80% <sup>4</sup>, and
Two or more VLs ≥ 1000 c/mL

taken two or more years after starting TLD regimen
or at least one VL ≥ 1000 c/mL and either

CD4 < 200 cells/mm3 or an opportunistic infection

If adherence still suboptimal <sup>4</sup>, or persistent low-level viraemia (2 or more consecutive VLs between 50 and 999 c/mL)



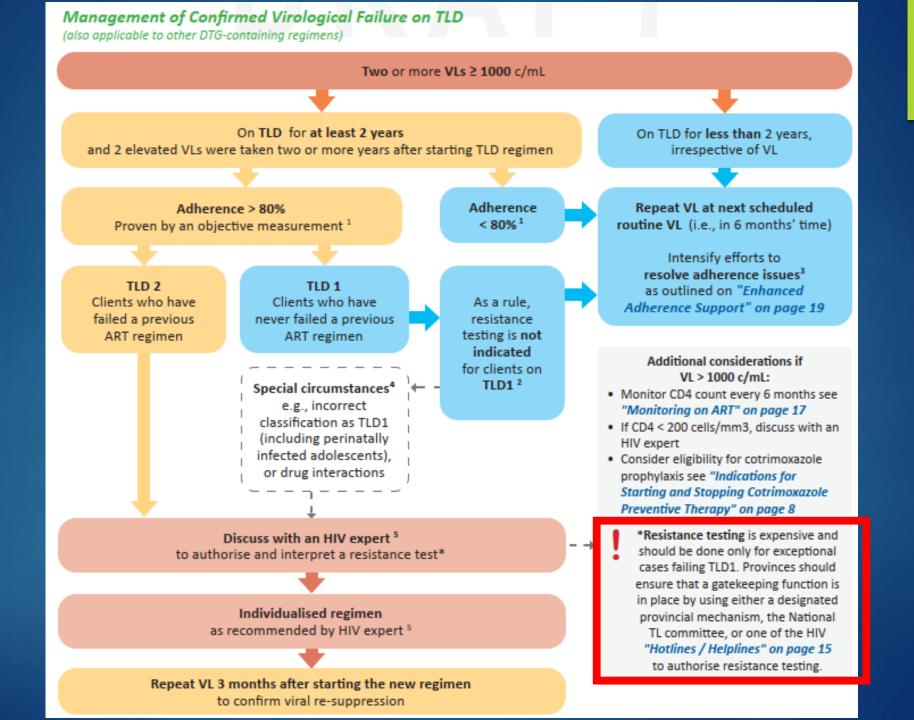
Repeat VL at next scheduled routine VL (i.e., in 6 months' time) Intensify efforts to resolve adherence issues <sup>2</sup>

### Management of Confirmed Virological Failure on TLD (also applicable to other DTG-containing regimens) Two or more VLs ≥ 1000 c/mL On TLD for at least 2 years On TLD for less than 2 years, and 2 elevated VLs were taken two or more years after starting TLD regimen irrespective of VL Adherence Repeat VL at next scheduled Adherence > 80% < 80% 1 routine VL (i.e., in 6 months' time) Proven by an objective measurement 1 Intensify efforts to resolve adherence issues<sup>3</sup> TLD 2 TLD 1 as outlined on "Enhanced Clients who have Clients who have As a rule, Adherence Support" on page 19 failed a previous never failed a previous resistance testing is not ART regimen ART regimen indicated Additional considerations if for clients on VL > 1000 c/mL: TLD1<sup>2</sup> Special circumstances . Monitor CD4 count every 6 months see e.g., incorrect "Monitoring on ART" on page 17 classification as TLD1 If CD4 < 200 cells/mm3, discuss with an</li> (including perinatally HIV expert infected adolescents), · Consider eligibility for cotrimoxazole or drug interactions prophylaxis see "Indications for Starting and Stopping Cotrimoxazole Preventive Therapy" on page 8 \*Resistance testing is expensive and Discuss with an HIV expert 5 should be done only for exceptional to authorise and interpret a resistance test\* cases failing TLD1. Provinces should ensure that a gatekeeping function is in place by using either a designated Individualised regimen provincial mechanism, the National as recommended by HIV expert 5 TL committee, or one of the HIV "Hotlines / Helplines" on page 15 to authorise resistance testing. Repeat VL 3 months after starting the new regimen to confirm viral re-suppression

# Footnote 4: Special circumstances that may warrant a resistance test for clients on TLD1 include

- Incorrect classification as TLD1 (clients who declare themselves as never having had ART before, but who have actually been exposed to ART and may have failed a regimen in the past)
- Perinatally infected adolescents: Unless a clearly documented drug history is available, perinatally infected adolescents should be classified as TLD2 due to the high likelihood of ART exposure and virological failure in the past
- Current or previous drug interactions with rifampicin, carbamazepine, phenytoin, phenobarbital, or the polyvalent cations may have resulted in the development of resistance. Drug interactions may also warrant an expert discussion and authorisation of a resistance test earlier than 2 years on the regimen.

In these types of exceptional circumstances, TLD1 clients with persistent virological failure despite confirmed good adherence may be discussed with an expert to authorise a resistance test on a case-by-case basis.



### **HELPLINES**



If in doubt about any aspect of viral load management or switching to second-line, contact one of the following resources:



National HIV & TB Health Care Worker Hotline: 0800 212 506



Right to Care Paediatric, Adolescent and Adult HIV Helpline: 082 352 6642



KZN Paediatric Hotline: 0800 006 603

## Right to Care HELPLINE

For nurses, doctors, pharmacists and other health care workers needing expert advice on all paediatric, adolescent and adult HIV and TB management.

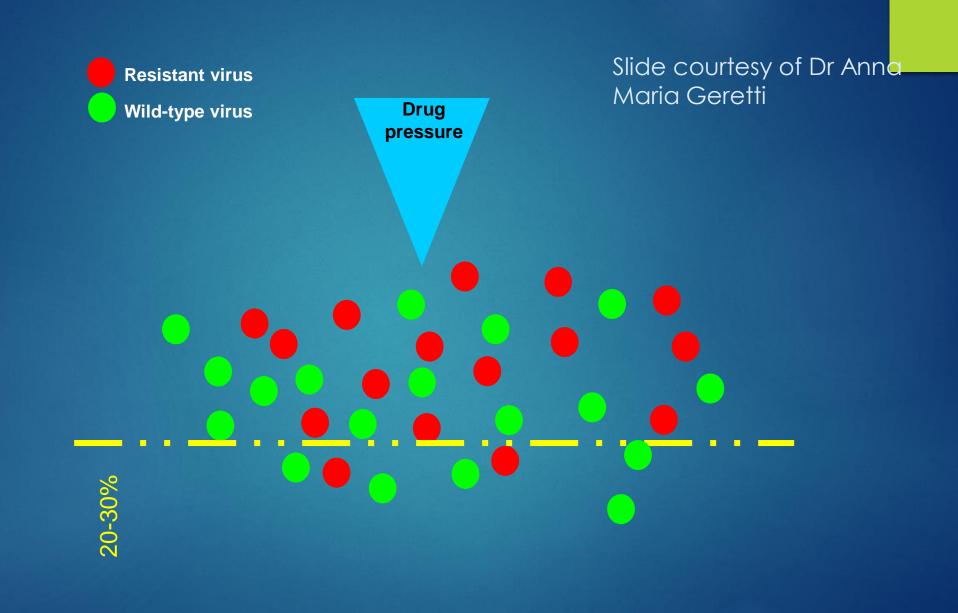
Call during office hours "please call me", sms or whatsapps may be sent and we can call you back.

HIV Helpline (adult and paediatric)



# How to do resistance testing

Patient must be adherent before doing resistance testing



### All Mutations Detected (HXB2 reference Sequence) Resistance mutations in bold

V35T, E36A, T39D, S48T, D123N, K173A, Q174K, D177E, T200A, Q207E, L214F, V245Q, E248D, I274V, R277K, Q278H, K281R, T286A, E291D, D324E, I329L, Q334N, G335D, R356K, G359T,

Reverse transcriptase I274V, R277K, Q278H, K281R, T2

T376S, T377L, T386I

Protease V3I, T12S, I15V, L19T, M36I, S37N, R41K, H69K, I93L

| Class         | Drug            | * <u>STAN</u> • v6.2.0 29/05/2012 |
|---------------|-----------------|-----------------------------------|
| NRTI          | Zidovudine      | S                                 |
|               | Didanosine      | S                                 |
|               | Stavudine       | S                                 |
|               | Lamivudine      | S                                 |
|               | Emtricitabine   | S                                 |
|               | Abacavir        | S                                 |
|               | Tenofovir       | S                                 |
| NNRTI         | Nevirapine      | S                                 |
|               | Efavirenz       | S                                 |
|               | Etravirine      | S                                 |
|               | Rilpivirine     | S                                 |
| PI/Boosted PI | Indinavir/r     | S                                 |
|               | Saquinavir/r    | S                                 |
|               | Nelfinavir      | S                                 |
|               | Fosamprenavir/r | S                                 |
|               | Lopinavir/r     | S                                 |
|               | Atazanavir/r    | S                                 |
|               | Tipranavir/r    | S                                 |
|               | Darunavir/r     | S                                 |

# How to do resistance testing

- Ask patient to take ARVs regularly for 1 month and then do resistance testing
- Call an expert to authorise and find out how to do the resistance test and again so they can help interpret the results
- ▶ If PI resistance apply to Third Line ART (TLART) Committee
  - ▶ TLART@health.gov.za
  - ▶ Third Line ART website: <a href="https://www.righttocare.org/what-we-do/third-line-art/">https://www.righttocare.org/what-we-do/third-line-art/</a>

# If there is DTG resistance

### Third Line ART:

- Likely DRV/r-based regimen
- Discuss with an expert

very

important things to remember

## For everyone with a VL>50



Do a thorough assessment of the cause of an elevated VL. Consider the possibility of:

- A. Adherence problems (see "Enhanced Adherence Support" on page 22)
- B. <u>Bugs</u> (Intercurrent infections)
- C. In-Correct ART dosage (see A ure 5 "Drug Dosing Chart" on page 34)
- D. Drug Interactions (see"Drug | ractions with DTG and Rifampicin-containing TB Treatment" on page 13)
- E. REsistance (if > 2 years on treent)

Implement interventions to re-suppress the VL, Luding Enhanced Adherence Support if indicated

Recommend condom use and contraception as appropriate

Two or more VLs ≥ 1000 c/mL

### Additional considerations if VL > 1000 c/mL:

- Monitor CD4 count every 6 months see
   "Monitoring on ART" on page 17
- If CD4 < 200 cells/mm3, discuss with an HIV expert
- Consider eligibility for cotrimoxazole prophylaxis see "Indications for Starting and Stopping Cotrimoxazole Preventive Therapy" on page 8

Individualised regimen as recommended by HIV expert 5

Repeat VL 3 months after starting the new regimen to confirm viral re-suppression

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provincial mechanism, the National TL committee, or one of the HIV "Hotlines / Helplines" on page 15 to authorise resistance testing.

# If CD4<200 provide AHD package of care

## Discuss with an expert



**Screening for Ols** 



**Prophylaxis:** 



Diagnosis and management of Ols

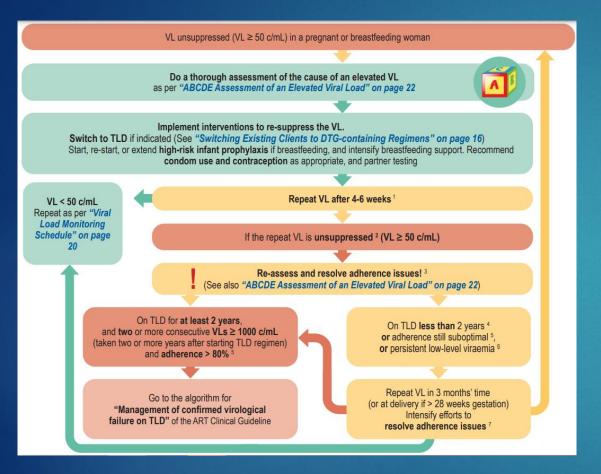


Risk of IRIS when starting or switching ART



Urgent help to achieve treatment optimisation and viral suppression

| Medical Indications to Defer ART  |   |  |  |  |
|---|---|--|--|--|
| Indication  | Action  |  |  |  |
| TB symptoms (cough, night sweats, fever, recent weight loss)  | Investigate symptomatic clients for TB before initiating ART. If TB is excluded, proceed with ART initiation and TB preventive therapy (after excluding contraindications to TPT). If TB is diagnosed, initiate TB treatment and defer ART. The timing of ART initiation will be determined by the site of TB infection and the client's CD4 cell count   |  |  |  |
| Diagnosis of drug-sensitive (DS) TB at a non-neurological site (e.g. pulmonary TB, abdominal TB, or TB lymphadenitis)   | <ul> <li>Defer ART initiation as follows:</li> <li>If CD4 &lt; 50 cells/μL − initiate ART within 2 weeks of starting TB treatment, when the client's symptoms are improving, and TB treatment is tolerated</li> <li>If CD4 ≥ 50 cells/μL − initiate ART 8 weeks after starting TB treatment</li> <li>In pregnant and breastfeeding women (PBFW) initiate ART within 2 weeks of starting TB treatment, when the client's symptoms are improving, and TB treatment is tolerated. Defer ART for 4-6 weeks if symptoms of meningitis are present. For further details, refer to the Family-Centered Transmission Prevention Guideline 2023</li> </ul> |  |  |  |
| Diagnosis of drug-resistant (DR) TB at a non-neurological site (e.g. pulmonary TB, abdominal TB, or TB lymphadenitis)   | Initiate ART after 2 weeks of TB treatment, when the client's symptoms are improving, and TB treatment is tolerated   |  |  |  |
| Diagnosis of DS-TB or DR-TB at a neurological site (e.g. TB meningitis or tuberculoma)  | Defer ART until 4-8 weeks after start of TB treatment   |  |  |  |
| Signs and symptoms of meningitis  | Investigate for meningitis before starting ART  |  |  |  |
| Cryptococcal antigen (CrAg) positive in the absence of symptoms or signs of meningitis and if lumbar puncture is (LP) negative for cryptococcal meningitis (CM) | No need to delay ART. ART can be started immediately.   |  |  |  |
| Confirmed cryptococcal meningitis   | Defer ART until 4-6 weeks of antifungal treatment has been completed  |  |  |  |
| Other acute illnesses e.g.  Pneumocystis jirovecii pneumonia (PJP) or bacterial pneumonia   | Defer ART for 1-2 weeks after commencing treatment for the infection  |  |  |  |
| Clinical symptoms or signs of liver disease   | Confirm liver injury using ALT and total bilirubin levels. ALT elevations > 120 IU/L with symptoms of hepatitis, and/or total serum bilirubin concentrations > 40 µmol/L are significant. Investigate and manage possible causes including TB, hepatitis B, drug-induced liver injury (DILI), or alcohol abuse  |  |  |  |



3

# Ask all WOCP with a high VL if they are pregnant or breastfeeding

Discuss with an expert

4

NHLS tools

High VL RfA (results for action)

Maternal VL eGK codes (electronic gatekeeping)

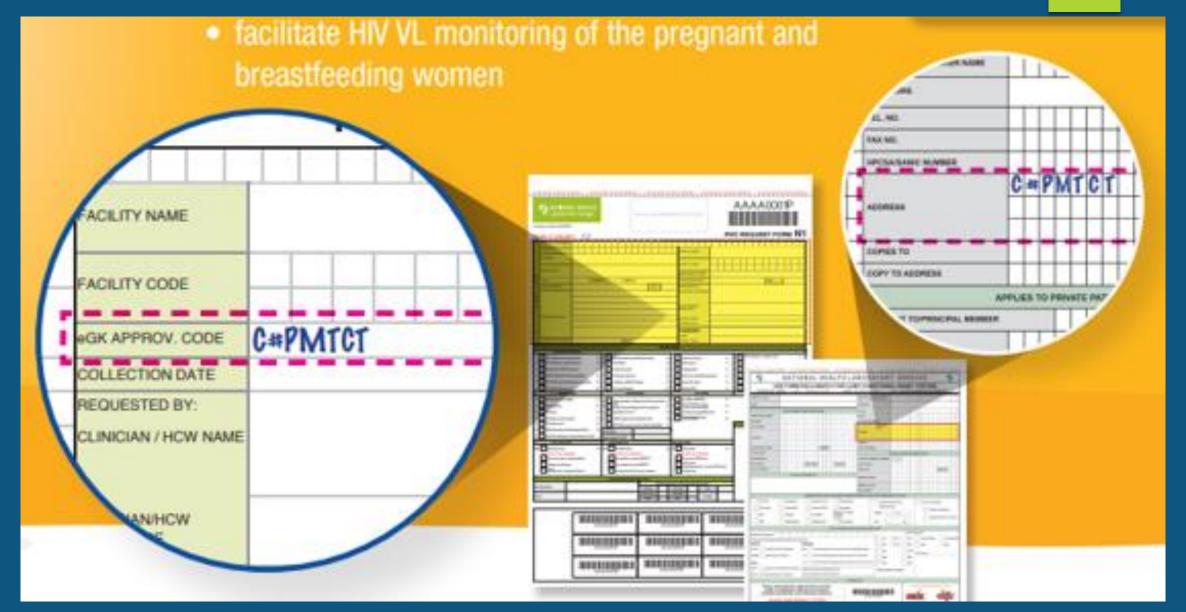
### EGK codes serve three functions:

Prevent sample rejection

2 Allow individual patients to be traced using the NHLS RfA reports

Monitor VL suppression rates at a program level

# C#PMTCT – pregnancy & breastfeeding C#DELIVERY – labour/delivery



# Summary

- Need to act on high VLs urgently
- Resistance develops quicker in NNRTIs than in PI's or DTG, therefore we approach them differently
- Aim to get most patients onto TLD
- Patients must be adherent to have accurate resistance test
- When in doubt call for help

### VL>50

On EFV/ NVP Or PI <2 years Address ABCDE
Review side effects
If VL>1000: Take CD4 count,
provide condoms and
contraception ask WOCP if
pregnant/bf

On LPV/r > 2 years

Change to TLD (regardless of VL)

Assess ABCDE & Call helpline
Resistance test if >80% adherent

On TLD2 > 2 yrs or drug interaction

Assess ABCDE & Call helpline
Resistance test if >80% adherent

20 year old female on TEE with a high VL. what will you do?

- ► CD4 count
- SRH counselling, condoms and contraception
- Assess ABCDE
- ► EAC and switch to AZT 3TC LPV/r
- ► EAC and switch to AZT 3TC DTG
- ► EAC and switch to TLD1
- ► EAC and switch to TLD2
- Call a helpline/ID consultant

36 year old male who was on TEE and was switched to TDF 3TC LPV/r in 2022 due to treatment failure. He now has a high VL. What will you do?

- ► CD4 count
- SRH counselling and condoms
- Assess ABCDE
- ► EAC and continue TDF 3TC LPV/r
- ► EAC and switch to AZT 3TC DTG
- ► EAC and resistance test
- ► EAC and switch to TLD2
- Call a helpline/ID consultant

17 year old female on AZT 3TC LPV/r since age 3, with multiple high VL over the years. She has not suppressed since 10 years old. She weighs 38kg and has a mild cough.

- ▶ CD4 count, Cr, FBC and differential
- Sputum GXP
- SRH counselling, condoms and contraception
- Assess ABCDE
- ► EAC and switch to TLD2 today
- ► EAC and HIV resistance test before switching to TLD2
- ► EAC and wait for blood results before switching to TLD2
- Call a helpline/ID consultant

Switching Existing Clients to DTG-containing Regimens
(Adults, adolescents or children who have never used a DTG-containing regimen in the past)

**VL-dependent regimen switches** Relevant to all clients who have been on PI-based regimens for more than two years: their VL result in the last 12 months

| will influence the decision of how and when to switch to a DTG-containing regimen          |  |   |   |  |
|--|--|---|---|--|
| VL<br>considerations   | Current Regimen  | Criteria for switch   | Regimen if change indicated   |  |
| VL < 1000 c/mL   | Any<br>LPV/r or ATV/r<br>regimen for<br>more than 2 years                            | Switch all to a DTG-containing regimen If VL in last 12 months was ≥ 50 c/mL, continue to switch same day, but do ABCDE assessment, provide EAC if needed, and repeat the VL after 3 months as per "The VL non-suppression algorithm" on page 19  | TLD  provided no renal dysfunction and age ≥ 10 yrs and weight ≥ 30 kg  If clients does not qualify for TDF  ABC¹/3TC/DTG |  |
| <sup>2</sup> Two or more VLs ≥ 1000 c/mL taken two or more years after starting PI regimen | Adult or adolescent on<br>regimen and adherence<br>less than 80% <sup>3</sup>        | Do not do a resistance test  These clients are unlikely to have PI resistance mutations. Rather switch to a more tolerable once daily 1 De regimen which is likely to support adherence. Manage as per "The VL non-suppression algorithm" on page 19  | TLD  provide an and age ≥ 10 yrs and weight ≥ 30 kg  If clients does not qualify for TDF  ABC¹/3TC/DTG                    |  |
|  | Adult or adolescent on<br>any<br>regimen and adherence<br>more than 80% <sup>3</sup> | Clients who meet the definition of confirmed virological failure and have confirmed adherence more than 80% may need a resistance test.  Discuss with an HIV expert <sup>4</sup> to authorise and interpret a resistance test.  Provide individualised regimen as recommended by HIV expert.  Repeat VL 3 months after the regimen change to confirm re-suppression, as per the "Management of Confirmed Virological Failure on TLD" on page 21 |   |  |
|  | Child < 10 years,<br>or weight < 30 kg<br>on any<br>LPV/r or ATV/r regimen           | These clients do not yet qualify for TLD and may require a resistance test.  Refer to algorithm "Switching children on PI-containing regimens to DTG-containing regimens" on page 16  |   |  |

55 year old male started TEE in 2014 and switched to TLD in 2019 with a suppressed VL.

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In 2020 and 2021 VL<50.
In 2022 VL: 1,2 mill.
In 2023 VL: 1.4 mill.
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- ► CD4 count
- Assess ABCDE
- SRH counselling and condoms
- ► EAC and switch to AZT 3TC LPV/r
- ► EAC and he needs third line ART
- ► EAC and consider HIV resistance test
- EAC and continue TLD
- Call a helpline/ID consultant

### Management of Confirmed Virological Failure on TLD (also applicable to other DTG-containing regimens) Two or more VLs ≥ 1000 c/mL On TLD for at least 2 years On TLD for less than 2 years, and 2 elevated VLs were taken two or more years after starting TLD regimen irrespective of VL Adherence Repeat VL at next scheduled Adherence > 80% < 80% 1 routine VL (i.e., in 6 months' time) Proven by an objective measurement 1 Intensify efforts to resolve adherence issues<sup>3</sup> TLD 2 TLD 1 as outlined on "Enhanced Clients who have Clients who have As a rule, Adherence Support" on page 19 failed a previous never failed a previous resistance testing is not ART regimen ART regimen indicated Additional considerations if for clients on VL > 1000 c/mL: TLD1<sup>2</sup> Special circumstances . Monitor CD4 count every 6 months see e.g., incorrect "Monitoring on ART" on page 17 classification as TLD1 If CD4 < 200 cells/mm3, discuss with an</li> (including perinatally HIV expert infected adolescents), · Consider eligibility for cotrimoxazole or drug interactions prophylaxis see "Indications for Starting and Stopping Cotrimoxazole Preventive Therapy" on page 8 \*Resistance testing is expensive and Discuss with an HIV expert 5 should be done only for exceptional to authorise and interpret a resistance test\* cases failing TLD1. Provinces should ensure that a gatekeeping function is in place by using either a designated Individualised regimen provincial mechanism, the National as recommended by HIV expert 5 TL committee, or one of the HIV "Hotlines / Helplines" on page 15 to authorise resistance testing. Repeat VL 3 months after starting the new regimen to confirm viral re-suppression

42 year old female was on TEE since 2017 then switched to AZT 3TC DTG due to treatment failure in 2020.

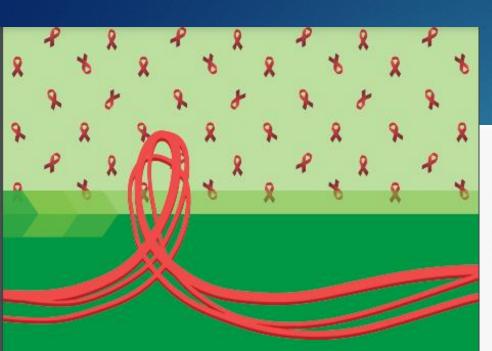
In 2021 VL: 2380

In 2022 VL: 4330

In 2023 VL: 12070

- ► CD4 count
- Assess ABCDE
- SRH counselling, condoms and contraception
- ► EAC and switch to TDF 3TC ATV/r
- ▶ EAC and he needs third line ART
- ► EAC and consider HIV resistance test
- EAC and switch to TLD
- Call a helpline/ID consultant

### Management of Confirmed Virological Failure on TLD (also applicable to other DTG-containing regimens) Two or more VLs ≥ 1000 c/mL On TLD for at least 2 years On TLD for less than 2 years, and 2 elevated VLs were taken two or more years after starting TLD regimen irrespective of VL Adherence Repeat VL at next scheduled Adherence > 80% < 80% 1 routine VL (i.e., in 6 months' time) Proven by an objective measurement 1 Intensify efforts to resolve adherence issues<sup>3</sup> TLD 2 TLD 1 as outlined on "Enhanced Clients who have Clients who have As a rule, Adherence Support" on page 19 failed a previous never failed a previous resistance testing is not ART regimen ART regimen indicated Additional considerations if for clients on VL > 1000 c/mL: TLD1<sup>2</sup> Special circumstances . Monitor CD4 count every 6 months see e.g., incorrect "Monitoring on ART" on page 17 classification as TLD1 If CD4 < 200 cells/mm3, discuss with an</li> (including perinatally HIV expert infected adolescents), · Consider eligibility for cotrimoxazole or drug interactions prophylaxis see "Indications for Starting and Stopping Cotrimoxazole Preventive Therapy" on page 8 \*Resistance testing is expensive and Discuss with an HIV expert 5 should be done only for exceptional to authorise and interpret a resistance test\* cases failing TLD1. Provinces should ensure that a gatekeeping function is in place by using either a designated Individualised regimen provincial mechanism, the National as recommended by HIV expert 5 TL committee, or one of the HIV "Hotlines / Helplines" on page 15 to authorise resistance testing. Repeat VL 3 months after starting the new regimen to confirm viral re-suppression



### 2023 ART Clinical Guidelines

for the Management of HIV in Adults, Pregnancy and Breastfeeding, Adolescents, Children, Infants and Neonates

June 2023 Version 4
Republic of South Africa National Department of Health



### Right to Care HELPLINE

For nurses, doctors, pharmacists and other health care workers needing expert advice on all paediatric, adolescent and adult HIV and TB management.

Call during office hours "please call me", sms or whatsapps may be sent and we can call you back.

HIV Helpline (adult and paediatric)









## **THANK YOU**

### DISCLAIMER

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