

# Using HIV-ASSIST to Guide ART Decision-Making

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## Disclosures

No conflicts of interest or relationships to disclose



### Outline

1. What is the HIV-ASSIST decision-support tool?

2. Use of HIV-ASSIST in different clinical scenarios



# HIV-ASSIST: Background



#### What is HIV-ASSIST?

- A free, online, evidence-based decision support tool for ART selection
- Developed by ID experts
- Uses multiple criteria decision analysis theory and evidence- and guideline-based effectiveness to rank different ART combinations
- Incorporates viral attributes, patient characteristics and preference, comorbidities, and co-medications
- Draws from DHHS and IAS-USA guidelines, clinical trials, Stanford Resistance Database,
   National HIV Curriculum, Liverpool HIV Drug Interactions, and a scientific advisory panel

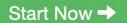


### Decision Support Tool for ART Selection

#### www.hivassist.com

# Welcome to HIV-ASSIST

HIV-ASSIST is a free, interactive, educational tool to inform clinical decision making for ARV selection



Take the tour >



Mutations	Mutations		Adherence	
Enter the patient's HIV mutations here, either with or (eg, M184V or 184V). You can use "INSERTION" or Separate mutations with commas. Example: 184V, 6  Viral Load  Suppressed (<50) for more than 6 months  Suppressed (<50) for less than 6 months or Low Level Viremia (<200)  Low (200 - 100,000)  High (100,000 - 500,000)  Very high (≥ 500,000)		Patients with pill aversion (prioritize smal Patients who prefer once daily dosing Patients with intermittent adherence Increase prioritization of at least 3 active Penalize regimens with IV/IM dosing Options for patients with poor adherence.  HLA-B5701 Positive (or unknown) Negative Select the patient's HLA-B5701 status (if known).		
Unknown Select the patient's HIV viral load (if known).				
Comorbidities, ARV Side Effects, or Pregnt Select the patient's comorbidities or side effects from pregnant.		Co-medications  Select the patient's current medications.		
Treatment History (Prior Failing Regimens)		Current Regimen		
Enter the patient's previous ART medications on whice regimen). Do not include the current regimen (if any)		Enter the patient's current regimen (if any).		



Mutations		Adherence	
Enter the patient's HIV mutations here, either with or (eg, M184V or 184V). You can use "INSERTION" or Separate mutations with commas. Example: 184V, 6		Patients with pill aversion (prioritize small Patients who prefer once daily dosing Patients with intermittent adherence Increase prioritization of at least 3 active Penalize regimens with IV/IM dosing Options for patients with poor adherence.	
Viral Load	CD4 Cell Count	HLA-B5701	Tropism
<ul> <li>Suppressed (&lt;50) for more than 6 months</li> <li>Suppressed (&lt;50) for less than 6 months or Low Level Viremia (&lt;200)</li> <li>Low (200 - 100,000)</li> <li>High (100,000 - 500,000)</li> <li>Very high (≥ 500,000)</li> <li>Unknown</li> <li>Select the patient's CD4 cell count (if known).</li> </ul>		<ul> <li>Positive (or unknown)</li> <li>Negative</li> <li>Select the patient's HLA-B5701 status (if known).</li> </ul>	<ul> <li>R5 virus</li> <li>X4 virus</li> <li>Dual Tropic virus</li> <li>Unknown</li> <li>Select the patient's HIV tropism (if known).</li> </ul>
Comorbidities, ARV Side Effects, or Pregnancy  Select the patient's comorbidities or side effects from current ARV medications, or if the patient is pregnant.		Co-medications  Select the patient's current medications.	
Treatment History (Prior Failing Regimens)		Current Regimen	
Enter the patient's previous ART medications on whi regimen). Do not include the current regimen (if any)		Enter the patient's current regimen (if any).	



Mutations		Adherence	
Enter the patient's HIV mutations here, either with or without the mutated amino acid but using uppercase		Patients with pill aversion (prioritize smaller pills) Patients who prefer once daily dosing Patients with intermittent adherence Increase prioritization of at least 3 active drugs Penalize regimens with IV/IM dosing Options for patients with poor adherence.	
Viral Load	CD4 Cell Count	HLA-B5701	Tropism
Suppressed (<50) for more than 6 months		Positive (or unknown)  Negative  Select the patient's HLA-B5701 status (if known).	<ul> <li>R5 virus</li> <li>X4 virus</li> <li>Dual Tropic virus</li> <li>Unknown</li> <li>Select the patient's HIV tropism (if known).</li> </ul>
Comorbidities, ARV Side Effects, or Pregnt Select the patient's comorbidities or side effects from pregnant.		Co-medications  Select the patient's current medications.	
Treatment History (Prior Failing Regimens	5)	Current Regimen	
Enter the patient's previous ART medications on whit regimen). Do not include the current regimen (if any)		Enter the patient's current regimen (if any).	



Mutations		Adherence	
		☐ Patients with pill aversion (prioritize smaller pills)	
		☐ Patients with intermittent adherence	
		☐ Increase prioritization of at least 3 active	drugs
Enter the nationals HIV mutations here, either with	or without the mutated amino acid but using uppercase	☐ Penalize regimens with IV/IM dosing	
(eg, M184V or 184V). You can use "INSERTION" ( Separate mutations with commas. Example: 184V	or "DELETION" as needed (eg, K67DELETION).	Options for patients with poor adherence.	
Viral Load	CD4 Cell Count	HLA-B5701	Tropism
<ul><li>Suppressed (&lt;50) for more than 6</li></ul>	<b>○</b> ≤ 50	O Positive (or unknown)	○ R5 virus
months	<b>○</b> ≤ 100	<ul><li>Negative</li></ul>	○ X4 virus
Suppressed (<50) for less than 6 month	S	Select the patient's HLA-B5701 status (if known).	O Dual Tropic virus
or Low Level Viremia (<200)  Cow (200 - 100,000)	> 200		<ul><li>Unknown</li></ul>
<ul><li>High (100,000 - 500,000)</li></ul>	○ Unknown		Select the patient's HIV tropism (if known).
○ Very high (≥ 500,000)	Select the patient's CD4 cell count (if known).		
○ Unknown			
Select the patient's HIV viral load (if known).			
Comorbidities, ARV Side Effects, or Pres	gnancy	Co-medications	
Select the patient's comorbidities or side effects from current ARV medications, or if the patient is pregnant.		Select the patient's current medications.	
Treatment History (Prior Failing Regimens)		Current Regimen	
Enter the patient's previous ART medications on w regimen). Do not include the current regimen (if ar	hich the patient had detectable viremia (i.e., failing y).	Enter the patient's current regimen (if any).	



Mutations		Adherence	
		<ul> <li>□ Patients with pill aversion (prioritize smaller pills)</li> <li>□ Patients who prefer once daily dosing</li> <li>□ Patients with intermittent adherence</li> </ul>	
Enter the patient's HIV mutations here, either with or (eg, M184V or 184V). You can use "INSERTION" or Separate mutations with commas. Example: 184V, 69	"DELETION" as needed (eg, K67DELETION).	☐ Increase prioritization of at least 3 active ☐ Penalize regimens with IV/IM dosing Options for patients with poor adherence.	drugs
Viral Load	CD4 Cell Count	HLA-B5701	Tropism
Suppressed (<50) for more than 6 months		Positive (or unknown)  Negative  Select the patient's HLA-B5701 status (if known).	<ul> <li>R5 virus</li> <li>X4 virus</li> <li>Dual Tropic virus</li> <li>Unknown</li> <li>Select the patient's HIV tropism (if known).</li> </ul>
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months  Suppressed (<50) for less than 6 months or Low Level Viremia (<200)  Low (200 - 100,000)  High (100,000 - 500,000)  Very high (≥ 500,000)  Unknown  Select the patient's HIV viral load (if known).	<ul> <li>≤ 100</li> <li>≤ 200</li> <li>&gt; 200</li> <li>Unknown</li> <li>Select the patient's CD4 cell count (if known).</li> </ul>	Negative  Select the patient's HLA-B5701 status (if known).	<ul> <li>X4 virus</li> <li>Dual Tropic virus</li> <li>Unknown</li> <li>Select the patient's HIV tropism (if known).</li> </ul>
Comorbidities, ARV Side Effects, or Pregnancy  Select the patient's comorbidities or side effects from current ARV medications, or if the patient is pregnant.  Treatment History (Prior Failing Regimens)		Co-medications  Select the patient's current medications.  Current Regimen	
Enter the patient's previous ART medications on whi regimen). Do not include the current regimen (if any)	ch the patient had detectable viremia (i.e., failing	Enter the patient's current regimen (if any).	



Mutations		Adherence	
Enter the patient's HIV mutations here, either with or (eg, M184V or 184V). You can use "INSERTION" or 'Separate mutations with commas. Example: 184V, 69	"DELETION" as needed (eg, K67DELETION).	Patients with pill aversion (prioritize small Patients who prefer once daily dosing Patients with intermittent adherence Increase prioritization of at least 3 active Penalize regimens with IV/IM dosing Options for patients with poor adherence.	
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Treatment History (Prior Failing Regimens)		Current Regimen	
Enter the patient's previous ART medications on which regimen). Do not include the current regimen (if any).		Enter the patient's current regimen (if any).	

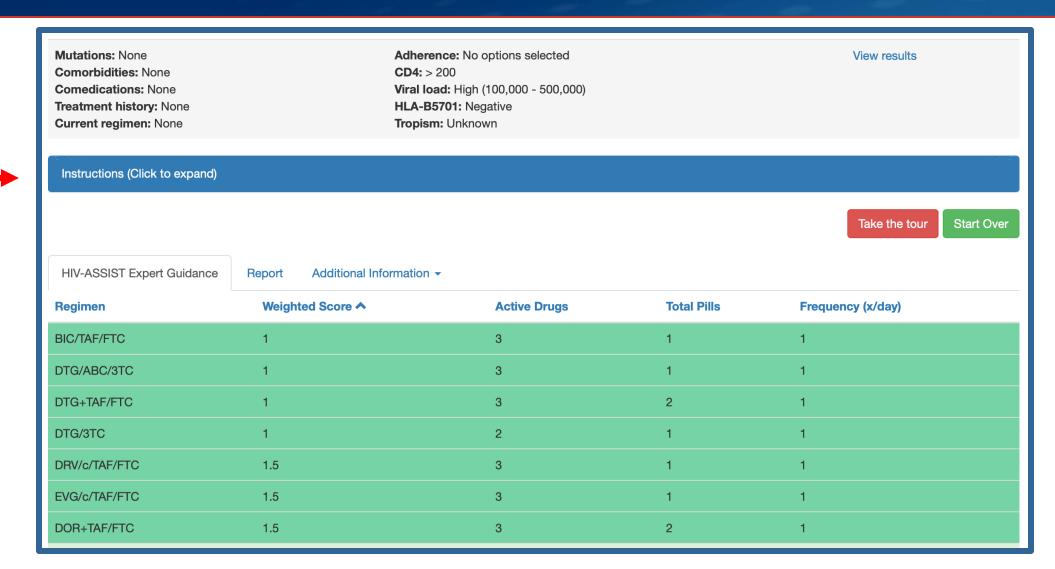


# HIV-ASSIST Inputs

3TC (Lamivudine/Epivir)	<ul><li>LPV/r (Lopinavir-ritonavir/Kaletra)</li></ul>	ATV (Atazanavir/Reyataz)
FTC (Emtricitabine/Emtriva)	FPV/r (Fosamprenavir-ritonavir/Lexiva and	□ DRV (Darunavir/Prezista)
ABC (Abacavir/Ziagen)	Norvir)	DRV/r (Darunavir-ritonavir/Prezista and
TAF (Tenofovir/Vemlidy)	TPV/r (Tipranavir-ritonavir/Aptivus and	Norvir)
TDF (Tenofovir/Viread)	Norvir)	□ DRV/c (Darunavir-cobicistat/Prezcobix)
AZT (Zidovudine/Generic)	SQV/r (Saquinavir-ritonavir/Invirase and	☐ RAL (Raltegravir/Isentress)
D4T (Stavudine/Zerit)	Norvir)	EVG/c (Elvitegravir-cobicistat/Vitekta)
DDI (Didanosine/Videx)	IDV/r (Indinavir-ritonavir/Crixivan and	□ DTG (Dolutegravir/Tivicay)
EFV (Efavirenz/Sustiva)	Norvir)	☐ BIC (Bictegravir/Biktarvy)
ETR (Etravirine/Intelence)	NFV (Nelfinavir/Viracept)	CAB (Cabotegravir/Cabenuva)
RPV (Rilpivirine/Edurant)	ATV/r (Atazanavir-ritonavir/Reyataz and	
NVP (Nevirapine/Viramune)	Norvir)	☐ IBA (Ibalizumab/Trogarzo)
DOR (Doravirine/Pifeltro)	ATV/c (Atazanavir-cobicistat/Evotaz)	☐ FOS (Fostemsavir/Rukobia)
Check any ARVs you would like to exclude dua ARVs from the results. ARVs that are in the curtion in the first that is your preferred regimen?		red ARVs are pre-selected for exclusion. This will exclude any regimens that include the checked
Select the ART regimen you are considering fo	or this patient.	



### HIV-ASSIST: Regimen Outputs



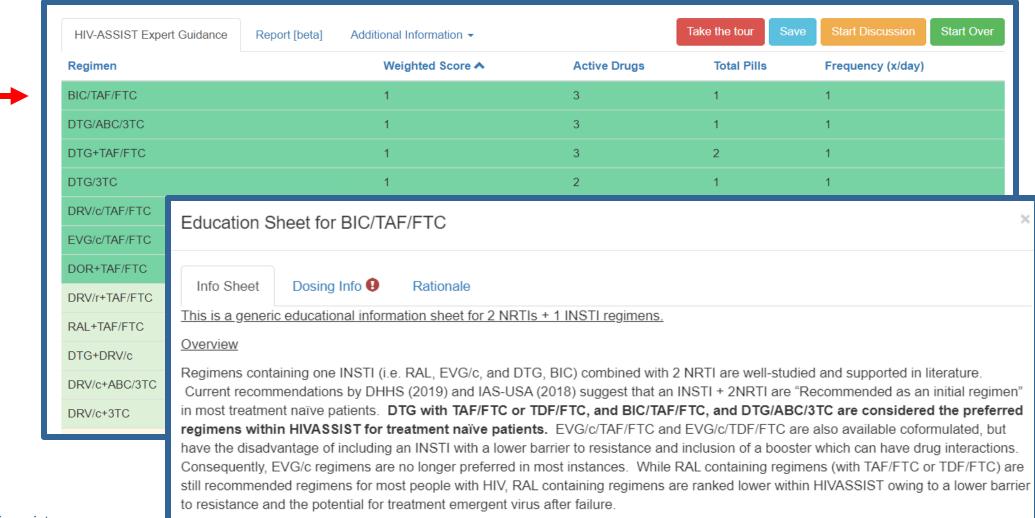


### Instructions on How to Interpret Outputs

Home / ARV Selection Tool / Results Mutations: None Adherence: No options selected View results Comorbidities: None CD4: > 200 Comedications: None Viral load: High (100,000 - 500,000) Treatment history: None HLA-B5701: Negative Current regimen: None Tropism: Unknown Instructions (Click to expand) HIV-ASSIST calculates a regimen's weighted score on a scale of 1 to 10, in which lower values represent preferred regimens based upon current IAS and DHHS quidelines, while factoring in treatment history, comorbidities, comedications, and other factors entered on the input page. The weighting system utilizes standardized utility weights based on current literature evidence and expert opinion. HIV-ASSIST is an educational tool and not a substitute for clinical judgement. Click on a regimen and 'Rationale' to see all steps leading to the presented weighted score. Please note that rankings would differ if if some factors were weighed more or less than those applied within HIV-ASSIST algorithms. HIV-ASSIST Score Efficacy (likelihood of viral suppression) Tolerability (side-effects and pill burden) Notes ≤ 1.5 Strong evidence Strong evidence Reserved for fully- or near fully- active regimens. 1.5 - 2.5 Strong or moderate evidence Strong or moderate evidence Includes regimens that started at better rank but were impacted by mutations, comorbidities, and drug interactions. 2.5 - 4.0Moderate evidence Moderate evidence As above. Additionally, most treatment-experienced patients will see regimens in this category. 4.0 - 6.0 Moderate to poor evidence Moderate or poor evidence As above. Additionally, most treatment-experienced patients will see regimens in this category. ≥ 6.0 Poor evidence Poor evidence As above. Additionally, these represent salvage regimens in patients with limited options. Signifies that there may be additional regimen warnings. Note: We do not show regimens for which there is evidence against usage.



### Double Click on Regimen for Educational Sheet





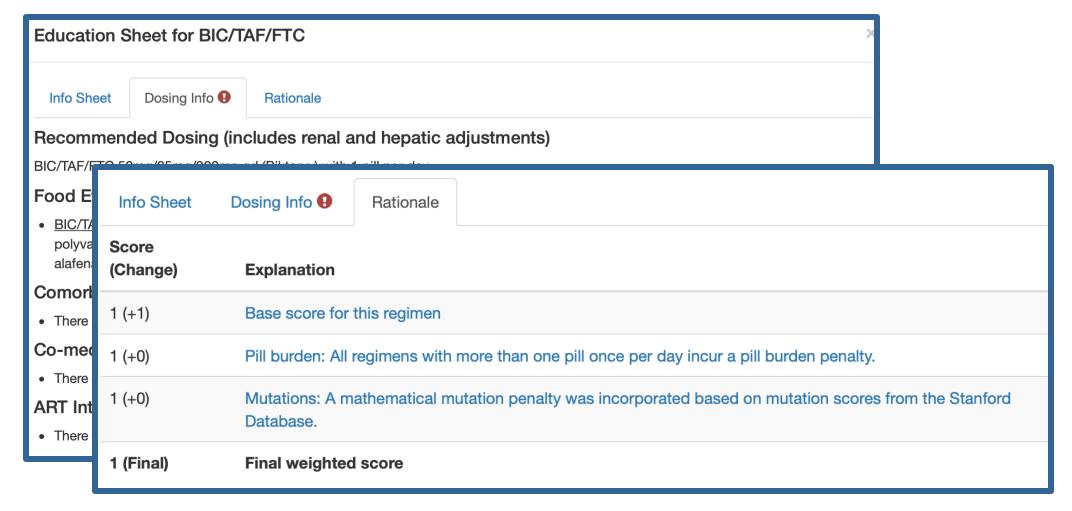
www.hivassist.com

# Education Sheet: Efficacy in Clinical Trials

TREATMENT	TREATMENT NAÏVE DTG or BIC			
Trial Name	Drugs Compared	Participants	Study Results	
SINGLE	ABC/3TC/ <b>DTG</b> vs. TDF/FTC/EFV	833 tx-naive	At week 48, the proportion of participants with an HIV-1 RNA level of less than 50 copies per milliliter was significantly higher in the ABC/3TC/DTG group than in the TDF/FTC/EVF group (88% vs. 81%). Was due primarily to discontinuations because of adverse events (2% in the ABC/3TC/DTG group and 10% in the TDF/FTC/EVF group). At week 144, ABC/3TC/DTG remained superior (71% vs 63% viral suppression).[9, 10]	
FLAMINGO	2 NRTIs plus DRV/r or <b>DTG</b>	484 tx-naive	At 48 weeks, DTG outperformed DRV/r (viral suppression 90% vs 83%, respectively).  Discontinuation due to adverse effects was higher in the DRV/r group than the DTG group (2% vs 4%, respectively), which contributed to the difference in the response rate. DTG continued to outperform DRV/r at 96 weeks (viral suppression 80% vs 66%)[11, 12]	
SPRING-2	2 NRTIs plus DTG or RAL	822 tx-naive	At 48 and 96 weeks, once-daily DTG was non-inferior to twice-daily RAL (88% vs 85% viral suppression at 48 weeks, and 81% vs 76% at 96 weeks), with a similar safety profile[13, 14]	
ARIA	ABC/3TC/ <b>DTG</b> vs. TDF/FTC+ATV/r	495 tx- naive women	At 48 weeks, ABC/3TC/DTG was superior in terms of virologic suppression (82% vs 71%). There were fewer virological nonresponses and fewer discontinuations due to adverse events in the ABC/3TC/DTG arm[15]	
Trial 1490	TAF/FTC/BIC vs. TAF/FTC/DTG	657 tx-naive	At week 96, HIV-1 RNA less than 50 copies per mL was achieved by 269 (84%) of 320 participants in the bictegravir group and 281 (86%) of 325 in the dolutegravir group (difference -2·3%, 95% CI -7·9 to 3·2), demonstrating non-inferiority of the bictegravir regimen compared with the dolutegravir regimen.[16]	
Trial 1489	TAF/FTC/ <b>BIC</b> vs. ABC/3TC/DTG	631 tx-naive	At week 96, bictegravir, emtricitabine, and tenofovir alafenamide was non-inferior to dolutegravir, abacavir, and lamivudine, with 276 (88%) of 314 participants in the bictegravir group versus 283 (90%) of 315 participants in the dolutegravir group achieving HIV-1 RNA less than 50 copies per mL (difference -1.9%; 95% CI -6.9 to 3.1).[17]	



## Education Sheet: Dosing and Rationale





#### HIV-ASSIST in the Literature

#### Development and validation of the tool<sup>1</sup>

- Cohort of 17 experienced providers reviewing 10 hypothetical cases
- 99% concordance for ARV-naïve patient cases
- 84-88% concordance for ART-experienced patient cases

#### Evaluating the concordance of the tool<sup>2</sup>

- Retrospective chart review of 106 patients at two clinics
- 100% concordance for ART-naïve patients
- 88-89% concordance for ART-experienced patients

#### Using HIV-ASSIST vs DHHS Guidelines<sup>3</sup>

- Randomized study of 118 medical trainees given HIV-ASSIST tool or DHHS guidelines to select ART for 10 hypothetical patient case scenarios
- Appropriate ART selections made 40% of the time with guidelines vs 90% of the time with HIV-ASSIST



# Using HIV-ASSIST in Different Clinical Scenarios



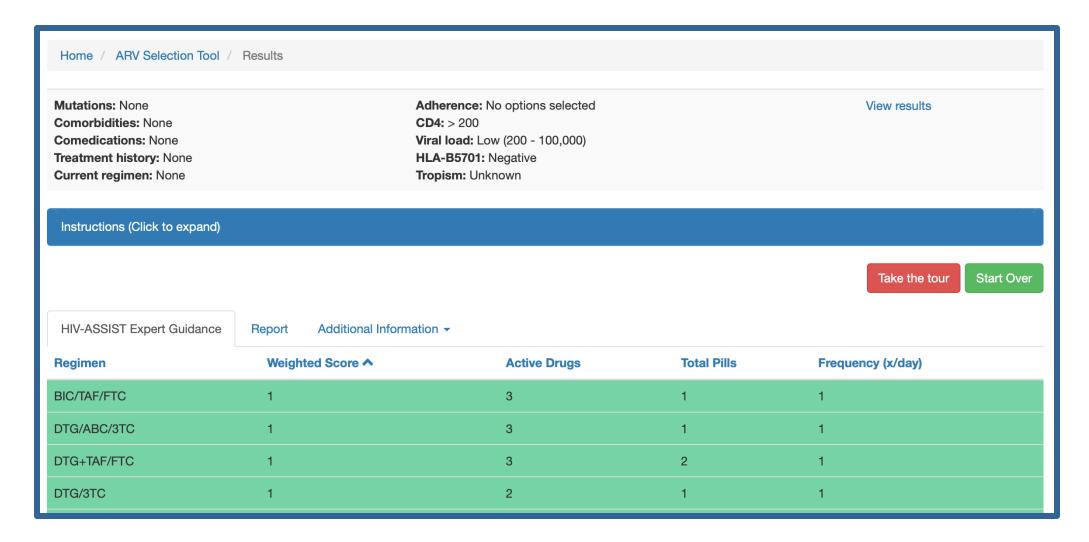
### Input: ART Start in a Newly Diagnosed PWH

A 26-year-old man presents with a new diagnosis of HIV. CD4 cell count is 360 cells/mm<sup>3</sup>. HIV-1 RNA is 34,000 copies/mL.

Mutations		Adherence	
Enter the patient's HIV mutations here, either with or without the mutated amino acid but using uppercase (eg, M184V or 184V). You can use "INSERTION" or "DELETION" as needed (eg, K67DELETION). Separate mutations with commas. Example: 184V, 65R		Patients with pill aversion (prioritize smaller pills)  Patients who prefer once daily dosing  Patients with intermittent adherence  Increase prioritization of at least 3 active drugs  Penalize regimens with IV/IM dosing  Options for patients with poor adherence.	
Viral Load  Suppressed (<50) for more than 6 months  Suppressed (<50) for less than 6 months or Low Level Viremia (<200)  Low (200 - 100,000)  High (100,000 - 500,000)  Very high (≥ 500,000)  Unknown  Select the patient's HIV viral load (if known).	CD4 Cell Count	HLA-B5701  Positive (or unknown)  Negative  Select the patient's HLA-B5701 status (if known).	Tropism  R5 virus  X4 virus  Dual Tropic virus  Unknown  Select the patient's HIV tropism (if known).
Comorbidities, ARV Side Effects, or Pregnancy  Select the patient's comorbidities or side effects from current ARV medications, or if the patient is pregnant.		Co-medications  Select the patient's current medications.	

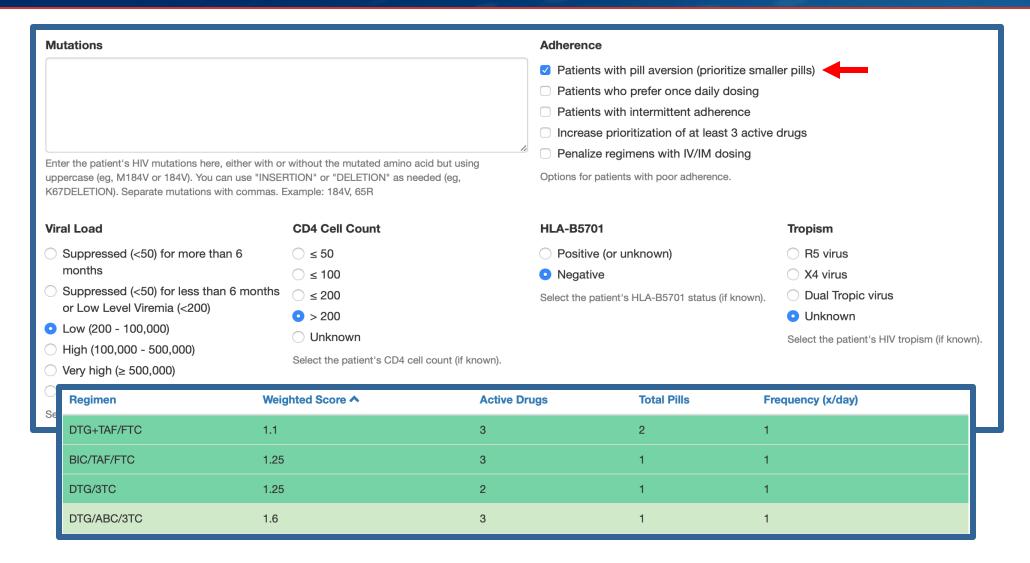


### Output: ART Start in a Newly Diagnosed PWH





### What if the patient has pill aversion?





# What if the baseline viral load is > 500,000?

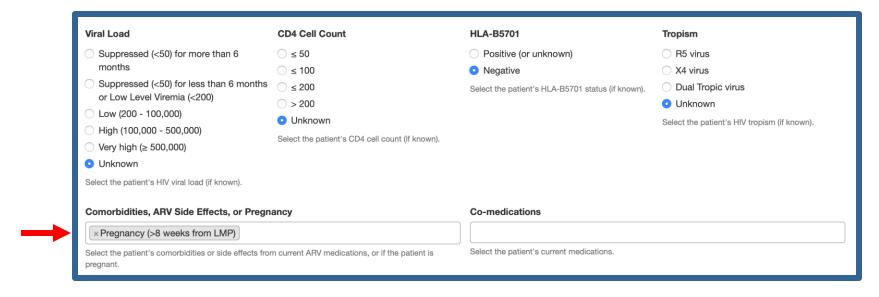
Viral Load	CD4 Cell Count	HLA-B5701	Tropism
<ul> <li>Suppressed (&lt;50) for more than 6 months</li> <li>Suppressed (&lt;50) for less than 6 months or Low Level Viremia (&lt;200)</li> <li>Low (200 - 100,000)</li> <li>High (100,000 - 500,000)</li> <li>Very high (≥ 500,000)</li> <li>Unknown</li> <li>Select the patient's HIV viral load (if known).</li> </ul>	<ul> <li>≤ 50</li> <li>≤ 100</li> <li>≤ 200</li> <li>&gt; 200</li> <li>Unknown</li> <li>Select the patient's CD4 cell count (if known).</li> </ul>	<ul> <li>Positive (or unknown)</li> <li>Negative</li> <li>Select the patient's HLA-B5701 status (if known).</li> </ul>	<ul> <li>R5 virus</li> <li>X4 virus</li> <li>Dual Tropic virus</li> <li>Unknown</li> <li>Select the patient's HIV tropism (if known).</li> </ul>

HIV-ASSIST Expert Guidance	Report Additional Information ▼			
Regimen	Weighted Score ◆	Active Drugs	Total Pills	Frequency (x/day)
BIC/TAF/FTC	1	3	1	1
DTG/ABC/3TC	1	3	1	1
DTG+TAF/FTC	1	3	2	1



### What if the person starting ART is pregnant?

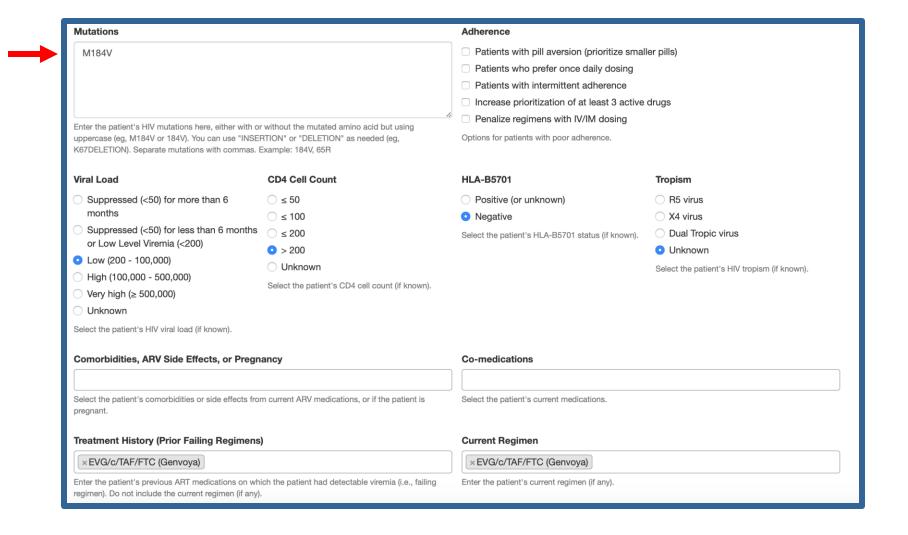
A 22-year-old woman is pregnant (16 weeks) and receives a positive screening HIV test at visit when she is establishing prenatal care.



Regimen	Weighted Score ◆	Active Drugs	Total Pills	Frequency (x/day)
DTG/ABC/3TC	1	3	1	1
DTG+TDF/FTC	1.15	3	2	1
DTG+TAF/FTC	1.2	3	2	1

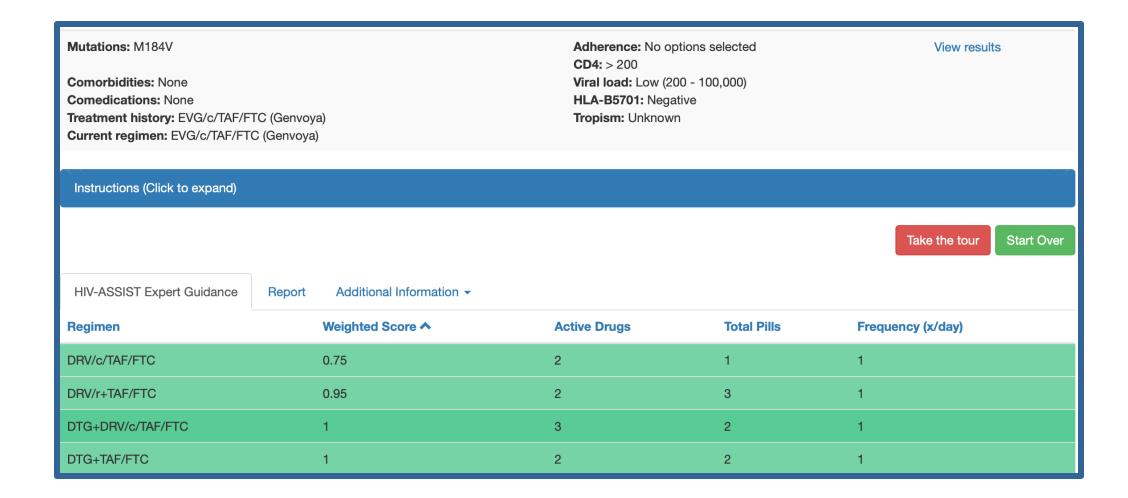


### What if the person has an M184V mutation?





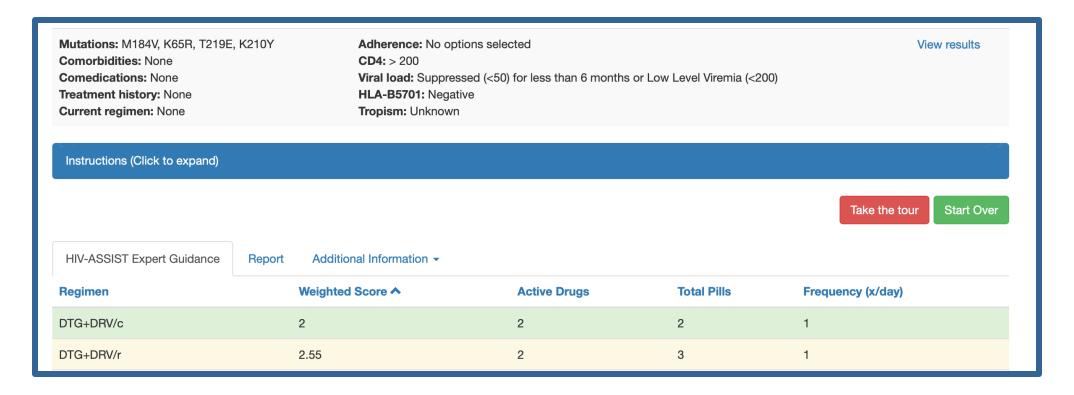
## Output: M184V that developed while on Genvoya®





### What if the person has multi-drug resistant HIV?

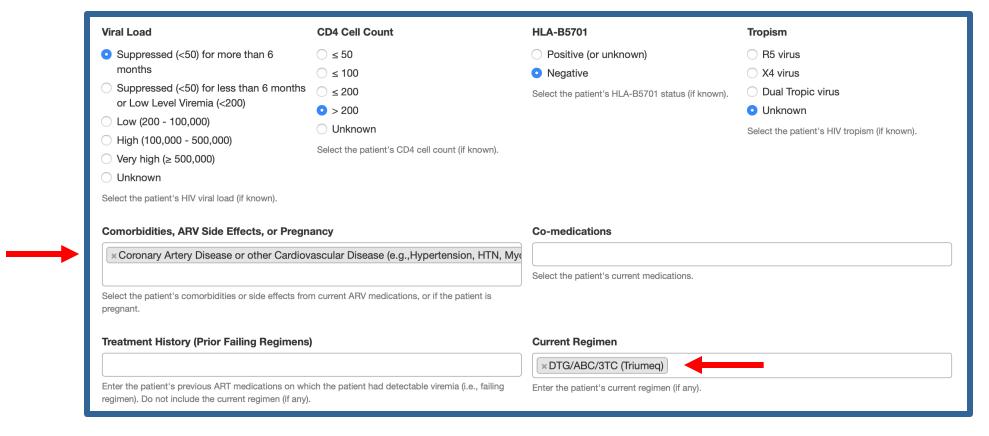
A 40-year-old with HIV (last CD4 350 cells/mm<sup>3</sup>, HIV-1 RNA undetectable) has M184V, K65R, T219E, and K210Y mutations.





#### **HIV-ASSIST & Comorbidities**

A 56-year-old man with CAD & HIV (last CD4 560 cells/mm<sup>3</sup> and HIV-1 RNA undetectable) on ABC/3TC/DTG is seen for follow-up.





# Output: Patient with heart disease on abacavir

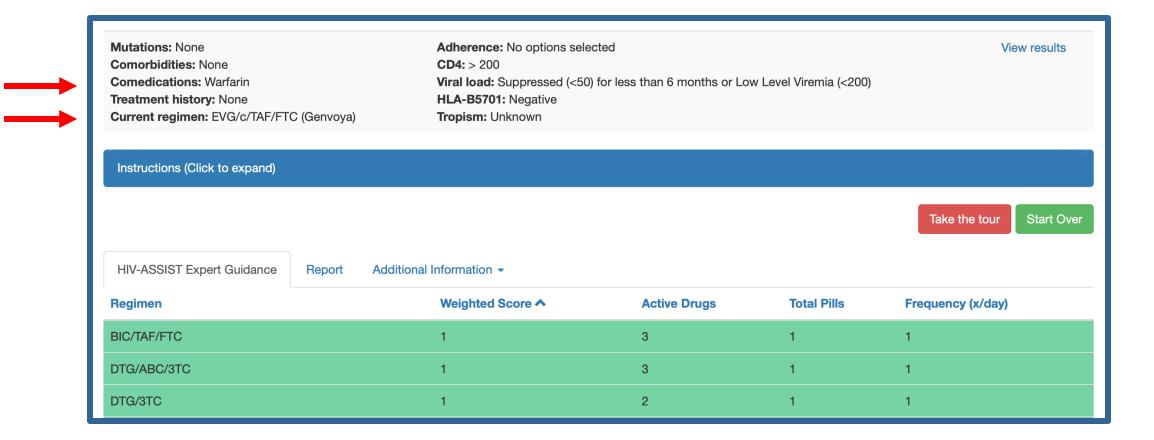
Regimen	Weighted Score ◆	Active Drugs	Total Pills	Frequency (x/day)
BIC/TAF/FTC	1	3	1	1
CAB/RPV	1	2	0	0.03
DTG/3TC	1	2	1	1

Regimen	Weighted Score ◆	Active Drugs	Total Pills	Frequency (x/day)
DRV/c+3TC	1.75	2	2	1
DTG/ABC/3TC [Current regimen]	2	3	1	1
DRV/r+DTG/3TC	2.4	3	3	1

Score (Change)	Explanation
1 (+1)	Suppressed viral load: Modified base score (for INSTI + NRTI +/- another ARV(s) with current viral suppression)
1 (+0)	Pill burden: All regimens with more than one pill once per day incur a pill burden penalty.
1 (+0)	Mutations: A mathematical mutation penalty was incorporated based on mutation scores from the Stanford Database.
2 (+1)	Comorbidities: This regimen incurred a penalty due to use of ABC in Coronary Artery Disease or other Cardiovascular Disease (e.g., Hypertension, HTN, Myocardial infarction).
2 (Final)	Final weighted score



### HIV-ASSIST & Drug-Drug Interactions



2.1



EVG/c/TAF/FTC [Current regimen]

#### Conclusions

- HIV-ASSIST is a helpful tool to aid in ART decision support
- The user can input as little or as much information as they would like
- Information comes from guidelines, clinical trials, Stanford Resistance Database, National HIV Curriculum, Liverpool HIV Drug Interactions, and a scientific advisory panel
- HIV/ID experts keep the tool up to date, and several new functionalities are on the way



# Acknowledgment

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