

# Switching from TDF to TAF: Missed Opportunities for Statin Use in HIV

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## Background

- People living with HIV (PLWH) have twice the risk for atherosclerotic cardiovascular disease (ASCVD) as the general population
- Increases in total and LDL cholesterol have been observed in PLWH switching from tenofovir disoproxil fumarate (TDF) to tenofovir alafenamide (TAF)<sup>1</sup>
- Current guidelines recommend initiating statin therapy in individuals with an elevated ASCVD risk (ASCVD ≥7.5%)²

# OBJECTIVE

To assess missed opportunities for statin therapy in PLWH switching from TDF- to TAF-containing antiretroviral therapy

# Methods

#### Study population OPERA cohort

- Prospectively captured, routine clinical data from electronic health records (84 clinics, 17 states, 1 US territory)
- » ~7% of PLWH in care in the US
- Inclusion criteria
- » HIV positive,  $\geq$ 18 years of age
- Switched from TDF to TAF between 5Nov2015 and 31Mar2018
- » ≥1 lipid measure on TDF ≤6 months prior to switch
- » ≥1 lipid measure ≥7 days after switch to TAF
- » No PEP, PrEP or clinical trial participation
- Observation period from TDF-to-TAF switch until the first censoring event:
- » TAF discontinuation
- $\gg \geq 12$  months without contact
- Death
- Study end (30Jun2018)

#### Lipid lowering agents

- Statins
- Prescription of atorvastatin, pitavastatin, pravastatin, rosuvastatin, or simvastatin
- With or without other lipid lowering agents
- Non-statins
- » Prescription of cholestyramine resin, ethyl eicosapentaenoic acid, ezetimibe, fenofibrate, gemfibrozil, niacin, Omega-3 acid ethyl esters, Omega-3 fatty acids
- Without statins

#### ASCVD risk score

- Derived from the Pooled Cohort Equations calculator<sup>2</sup>
- Predicts the risk of developing a first ASCVD event (i.e. nonfatal myocardial infarction, coronary heart disease death, or fatal or nonfatal stroke) over 10 years among people free from ASCVD
- Based on sex, age, race, total cholesterol, HDL, systolic blood pressure, hypertension treatment, diabetes and smoking status
- Imputed using limit value if age <40 or >79, systolic BP <90 or >200 mmHg, HDL <20 or</li> >100 mg/dL, total cholesterol <130 or >320 mg/dL

#### Analyses

- Proportion of PLWH prescribed statins pre- and post-switch
- Stratified by ASCVD risk

### Results Figure 1. Demographic and clinical characteristics of PLWH switching from TDF to TAF (N=6,451)



#### Figure 3. Risk of ASCVD in PLWH switching from TDF to TAF with an ASCVD<sup>+</sup> risk score



<sup>+</sup>All factors included in the ASCVD score calculation were measured within 6 months before the lipid pane

### No lipid-lowering agents\* Non-statins\* Statins\*† On TDF n=376 (6%) n=971 (15%) n=5104 (79%) 14% 79% 93% Non-statins<sup>‡</sup> No lipid-lowering agents<sup>‡</sup> Statins<sup>†‡</sup> On TAF n=1696 (26%) n=4132 (64%) n=623 (10%) \* Lipid lowering agent use assessed at time of lipid panel on TDF

† Statins used with or without other lipid lowering agents First lipid lowering agent use on/after first lipid panel on TAF

### Figure 4. Statin use by ASCVD\* risk in PLWH switching from TDF to TAF

Figure 2. Lipid-lowering agent use before and after switch (N=6,451)



\*All factors included in the ASCVD score calculation were measured within 6 months before the lipid panel

<sup>+</sup>Statin prescription at or after the last lipid panel on TDF <sup>‡</sup>Statin prescription at or after the first lipid panel on TAF



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### Discussion

- Changes in regimens represent an opportunity for healthcare providers to assess health markers and address clinical concerns
- A third of PLWH had an elevated ASCVD risk at the time of switch from TDF to TAF
- Failure to initiate statins in PLWH with an ASCVD  $\geq$ 7.5% at switch represents a missed opportunity for statin initiation
- » 69% had no statin prescription before switch
- » 59% had no statin prescription after switch
- Considerable missed opportunity given that the Pooled Cohort Equations calculator may underestimate risk among PLWH due to non-traditional risk factors driving ASCVD<sup>3</sup>
- Comparable results among PLWH who did not require ASCVD score imputation (data not shown)

# **KEY FINDINGS**

- Only an additional ~10% of PLWH with elevated ASCVD risk initiated statins after switching from TDF to TAF
- Considerable missed opportunities to reduce risk of cardiovascular disease in this at-risk population

### References

- 1. Mallon P, Brunet L, Fusco J, et al. Changes in Lipids After a Direct Switch from TDF to TAF. CROI Conference on Retroviruses and Opportunistic Infections. Seattle, WA, 2019.
- 2. Stone NJ, Robinson JG, Lichtenstein AH, et al. 2013 ACC/AHA Guideline on the Treatment of Blood Cholesterol to Reduce Atherosclerotic Cardiovascular Risk in Adults: A Report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines. Journal of the American College of Cardiology 2014; 63(25, Part B): 2889-934.
- 3. Zanni MV, Fitch KV, Feldpausch M, et al. 2013 American College of Cardiology/American Heart Association and 2004 Adult Treatment Panel III cholesterol guidelines applied to HIVinfected patients with/without subclinical high-risk coronary plaque. AIDS (London, England) 2014; 28(14): 2061-70.

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