

Single Tablet Regimens do not Necessarily Translate into More Durable HIV Treatments

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Background

In recent studies, Single Treatment Regimen (STR) have been associated with some better clinical outcomes:

- ↑ adherence
- ↓ virological failure
- ↓ hospitalization
- ↓ cost of health related problems

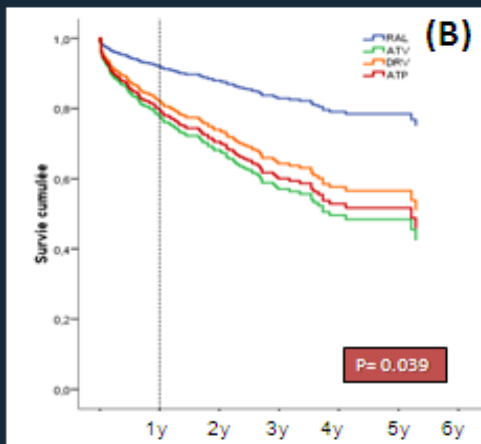
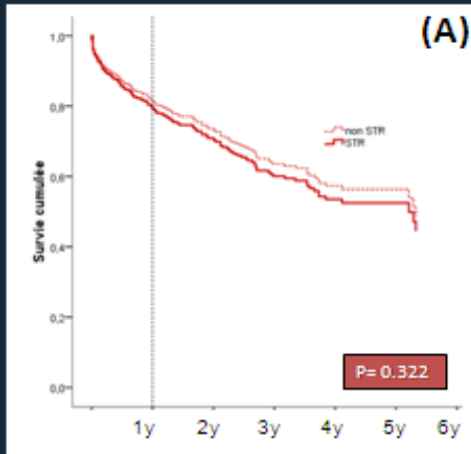
Methods

- Clinic-based, observational, retrospective study
- Inclusion: all patients starting any of the 4 recommended 1st regimens since 2007 : **TDF/FTC/EFV** as an STR, or any of the following non-STR: **2NRTI + either RAL, ATV/r, or DRV/r**
- Exclusion: switches for simplification & patients in a vaccine trial
- Primary endpoint: time until discontinuation of the 1st regimen (backbone change allowed)
- Secondary endpoint: time to loss of virological response at any time after start of 1st regimen. Switch allowed, D/C= failure
- Statistical analyses: Cox proportional hazards models. Controlling for: age, gender, baseline CD4, baseline HIV-RNA, HIV risk factor and calendar year of treatment initiation

RESULT-1: Durability

Table 3. Predictors of discontinuation of the first-line regimen

Parameters in the model	Hazard Ration (95% CI)	
	Simple Model	Adjusted Model
non-STR (vs STR)	0.86 (0.64-1.17)	
Third agent: ATP	<i>Ref</i>	<i>Ref</i>
RAL	0.45 (0.25-0.82)	0.36 (0.18-0.72)
ATV	0.96 (0.69-1.35)	1.21 (0.80-1.81)
DRV	0.99 (0.65-1.51)	0.99 (0.62-1.59)
Year of ARV initiation	1.13 (1.01-1.25)	1.21 (1.06-1.38)
IDU	1.66 (0.99-2.77)	2.14 (1.18-3.86)
Age	0.99 (0.97-1.01)	0.99 (0.97-1.01)
Men	0.87 (0.51-1.47)	0.90 (0.50-1.62)
VL@ARV1 > 100K	1.31 (0.97-1.78)	1.40 (1.03-1.91)



Durability of the first line regimen
 (A) stratified by STR vs non-STR
 (B) stratified by the 3rd ARV used

Univariate and multivariable Cox proportional hazards model of predictors of stopping first-line regimen

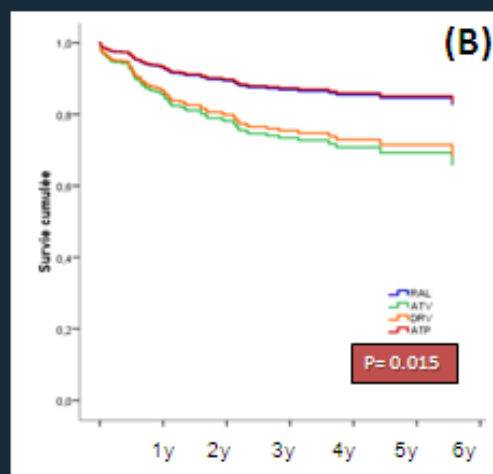
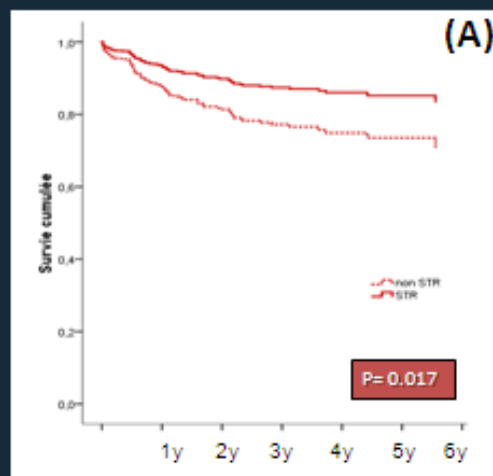
Main reason for D/C: Side effects & toxicity

Figure 2. Proportion of patients continuing the first-line regimen

RESULT-2: Virologic control

Table 4. Predictors of Loss of Virological Response (LOVR)

Figure 3. Proportion of patients maintaining virological response



Time to loss of virological response
(A) stratified by STR vs non-STR
(B) stratified by the 3rd ARV used

Parameters in the model	Hazard Ration (95% CI)	
	Simple Model	Adjusted Model
non-STR (vs STR)	1.85 (1.15-2.97)	
Third agent: ATP	<i>Ref</i>	<i>Ref</i>
RAL	1.11 (0.53-2.36)	1.04 (0.45-2.37)
ATV	2.03 (1.22-3.37)	2.28 (1.27-4.12)
DRV	2.10 (1.16-3.80)	2.09 (1.09-3.99)
Year of ARV initiation	1.02 (0.88-1.17)	1.12 (0.95-1.32)
IDU	2.70 (1.50-4.84)	2.57 (1.31-5.09)
Age	0.99 (0.98-1.02)	0.99 (0.97-1.01)
Men	0.64 (0.34-1.19)	0.79 (0.39-1.59)
VL@ARV1 > 100K	1.28 (0.85-1.93)	1.40 (0.92-2.15)

Univariate and multivariable Cox proportional hazards model of predictors of time to loss of Virologic response or treatment interruption

Conclusion

- STR does not provide a more durable 1st line treatment
- Even with a higher number of pills and multiple doses, a **2NRTI + RAL** regimen is **more durable** than ATP (STR) or other combinations (either ATV/r or DRV/r)
- The main reason for 1st line discontinuation/switch remain the adverse drug effects, very few virological failure were observed
- However, Initiating ARV treatment with an **STR** or a **RAL-based treatment** seems to provide a **longer virological control** compared to other non-STR regimens studied