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Implications of the human immunodeficiency virus test and treat strategy on antiretroviral treatment uptake and retention outcomes in Cameroon

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ABSTRACT

Background: Success of the human immunodeficiency virus (HIV) test-and-treat (T&T) strategy requires high antiretroviral (ART) uptake and retention. However, low ART uptake and retention continue to be reported in ART programs. This study assessed ART uptake and retention outcomes of the HIV T&T strategy in three HIV clinics in Cameroon.

Methods: A retrospective chart review was done for 423 patients who initiated HIV care within a period of three months prior to the implementation of the HIV T&T strategy, and for another 423 patients who initiated HIV care within a three-month period following the HIV T&T strategy implementation. For each group, sociodemographic, ART uptake and retention data were collected. Chi square and Student T tests were used to test for differences proportions and means between the two groups at p<0.05 and 95% confidence interval.

Results: The mean ages (years) in the pre-T&T and the T&T groups were 39.73 and 39.72, and the proportion of female were 65.85% and 65.08% respectively. ART uptake proportion was higher amongst those enrolled under the T&T strategy (98.08% vs 95.39%, p=0.02). A greater proportion of the patients in the T&T group initiated ART within 2 weeks following HIV diagnosis (55.84% vs 48.17%, p=0.03). However, ART retention at 24th month was lower in the T&T group (78.83% vs. 85.79%, p=0.01).

Conclusions: The findings suggest that the T&T strategy is associated with higher ART uptake, earlier ART initiation, and lower ART retention. This underscores a need for strategies to improve ART retention under the HIV T&T guidelines.

Keywords: Human immunodeficiency virus test and treat strategy, Antiretroviral treatment uptake, Antiretroviral treatment retention, Cameroon

INTRODUCTION

The human immunodeficiency virus (HIV) continues to be a major global public health problem. According to the World Health Organisation, an estimated 36.9 million people live with HIV, with over one million people dying of AIDS-related illnesses, and 1.8 million people newly

infected with HIV. The vast majority of this number live in low and middle income countries where the health systems are weak and unable to effectively control the epidemic.1

Several strategies have been adopted and implemented to control the HIV/AIDS pandemic with mixed results and progress in decreasing new HIV infections among adults

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has been slow in recent years. Since 2010, the annual number of new HIV infections among adults has remained around 1.9 million.² In 2015, the WHO recommended the HIV test and treat strategy as a global approach for the control of the HIV/AIDS pandemic. The strategy posits that expanded testing and earlier treatment of HIV infection could markedly decrease ongoing HIV transmission, stemming the HIV epidemic.³ The approach aims at increasing antiretroviral therapy (ART) coverage in order to attain community viral suppression through testing everyone for HIV, placing those tested HIV positive on ART irrespective of CD4 count or clinical criteria, and maintaining HIV positive people on ART in order to achieve sustained viral suppression. The scientific basis for this approach is supported by evidence that early initiation of ART is associated with better treatment outcome and people on highly active antiretroviral therapy (HAART) are less likely to transmit the virus to others due to viral suppression. 4-6 The overall goal of the universal test and treat (T&T) strategy is to end the AIDS epidemic as a public health threat by 2030, a target included in the 2030 agenda for sustainable development adopted by the United Nations General Assembly in September 2015. As new infections and HIV-related deaths are still unacceptably high, the immediate challenge is to reach the Fast-Track 2020 targets of reducing the number of people acquiring HIV to fewer than 500 000 and reducing the number of people dying from HIV-related causes to fewer than 500 000. Based on current estimates, this provides an opportunity to prevent over 1.6 million new infections and 600 000 deaths per year.³ In line with this WHO recommended approach to end the HIV/AIDS epidemic, the Joint United Nations Program on HIV/AIDS (UNAIDS) set the 2020 fast track 90-90-90 targets.⁷

Cameroon is one of the thirty-five countries accounting for more than 90% of the people becoming newly infected with HIV and designated Fast-Track countries requiring intensified action.³ With the ambition to accelerate antiretroviral therapy (ART) coverage as well as to reach the global objective of eliminating HIV by 2030 under the 90-90-90 response strategy, the Cameroon Ministry of Health, in May 2016, implemented the WHO HIV T&T strategy in all health facilities providing HIV services within the national territory. The HIV T&T strategy has been associated with mixed outcomes. While it has been associated with improved ART uptake and retention in some contexts, it has also showed lower ART retention in others, and there is paucity in literature on the HIV T&T outcomes in Cameroon.⁸⁻¹¹ ART uptake and retention outcomes of the HIV T&T strategy, both representing important prerequisites for the success of the HIV T&T strategy, have not been extensively described in Cameroon. This study aimed at assessing ART uptake and retention outcomes of the HIV universal teat and treat strategy in three HIV clinics in Cameroon.

METHODS

Ethical considerations

An ethical clearance prior to the implementation of this study was obtained from the Faculty of Health Science Institutional Review Board of the University of Buea. 12 The study was implemented within the institutional framework of the Cameroon International epidemiology Database to Evaluate AIDS (IeDEA) study approved by the National Ethics Committee of the Cameroon's Ministry of Health.

Study design and data collection procedure

This is a comparative retrospective study in which a 24month longitudinal medical chart review was done for patients who enrolled in HIV care within a three-month period preceding the implementation of the HIV T&T guidelines and another group of patients who enrolled within another three-month period following the implementation of the guidelines in three HIV clinics in Cameroon. The HIV T&T strategy was implemented in Cameroon in May 2016. The three preceding months included February to April, 2016 and the three months following the implementation of the strategy included July to September 2016. The months of May and June were considered the transition period and were thus excluded. Medical records were reviewed chronologically and a structured form was used to abstract data on; Sociodemographic characteristics, HIV related baseline and follow up outcomes, including ART retention.

Study sites

The study included all the three Cameroon IeDEA HIV clinics, located in three different regions of Cameroon. The clinics included: the Bamenda Regional Hospital (BRH) in the North West Region, the Limbe Regional Hospital (LRH) in the South West Region, and the Yaoundé Jamot Hospital (YJH) in the Centre Region of Cameroon. The hospitals are all tertiary level, government owned facilities, with specialized HIV clinics and provided care to a total of about 12,500 HIV positive clients; BRH (5000), LRH (4000) and YJH (3500). Each of the HIV clinics had at least a medical doctor in charge of the treatment center, and other staff including nurses, pharmacy attendants, counsellors and psychosocial workers.

Study population, inclusion and exclusion criteria

This study included the clinic records of HIV positive adult patients who were enrolled in care in the study clinics, and felt within the study inclusion criteria. Patients whose medical files could not be traced were not included in the study. Patients transferred out at the time of enrolment in care at the clinic were excluded. Patients transferred in during the study recruitment period were not included.

Data collection

Two lists of clinic codes of subjects enrolled in HIV care during the study reporting periods, representing the pre T&T and the T&T groups, were generated chronologically from the respective clinics' enrolment registers, according to the date of first enrolment in care in the clinic. Using the generated lists for each group, the

subjects' medical records (medical charts, ART registers, ART dispensation registers) were sorted and sociodemographic, clinical, ART uptake and retention data were abstracted. In total, data was abstracted from 846 of 890 medical records reviewed. The number of records included per clinic was 252, 227 and 367 from Bamenda, Limbe and Jamot respectively. The data collection flow is schematically presented in the Figure 1.

Patients who initiated care in study clinics during study reporting period n=890

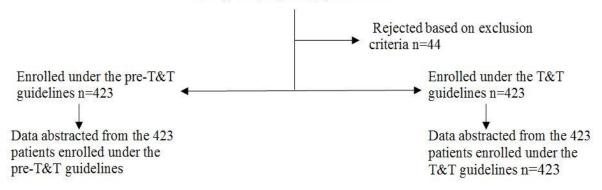


Figure 1: Data collection flow chart.

Data handling and statistical analysis

Data from de-identified paper forms were progressively entered in an Epi Info database by a trained data clerk using a laptop. Data cleaning was done both locally and centrally depending on the source of the data problem. Data cleaning rules and cut-off ranges were devised and provided to all three sites. At the central level, abstracted medical records were reviewed and double data entry was done by the study investigator to control for data quality. The paper forms were safeguarded in a locked cupboard after entry and the database backed up in an external hard drive.

The data was analyzed using Epi Info version 7 software. Means and standard deviations were calculated for continuous variables while frequency distributions calculated for categorical variables. Comparison of proportions and means were done using Chi square and the Student T-test respectively.

The study outcome variables were defined as follows:

ART retention

Proportion of subjects known to be accessed ART in the study clinics during the specific study follow up periods (6, 12 and 24 months).

ART uptake

Proportion of subjects enrolled in HIV care and initiated ART during the study reporting period.

Lost to follow up

Proportion of patients who failed to return for drug pickup after three consecutive months following the last expected pharmacy appointment date, and were not reported as deceased or transferred out.

Transferred out

Proportion of patients with clinic level documented evidence of being transferred from the study clinic to another HIV clinic.

Deceased

Proportion of patients with clinic level documented evidence of being deceased.

RESULTS

Baseline socio-demographic characteristics of the patients

A total of 846 records were analysed, constituting information from 423 files from participants in the pre T&T and 423 participants in the T&T groups. The mean ages (years) of the pre-T&T and the T&T groups were 39.72 ± 10.72 and 39.72 ± 11.10 . The proportion of males enrolled in the pre T&T and the T&T group were similar (34.14% vs 34.92%, p=0.81).

The proportions of patients with no formal and primary education were similar in both groups (7.6 vs. 7.11; 35.60

vs. 36.03). However, the proportion with secondary education was higher in the T&T group (48.77 vs. 43.48). More than two thirds of the participants in the pre T&T and the T&T group had a monthly income of less than 50,000 XAF (\$90), and the proportion was significantly higher in the T&T group (79.46 vs. 73,64, p=0.03). Unemployment proportion was slightly higher in the group enrolled under the T&T strategy (36.52% vs 31.41%, p=0.21).

Regarding the time spent traveling between patients' home and HIV clinic, more than half of subjects in both groups lived less than 30 minutes from their respective HIV clinics. However, the proportion was greater in the group under the T&T strategy (55.90 vs. 51.53, p=0.51).

Regarding substance use, the proportions of patients who smoked in the pre T&T and T&T groups were 15.27% and 13.13%. Alcohol intake proportions were respectively 63.04% and 61.16%. Most of those who consumed alcohol in both groups were of the male sex (78.89% and 71.90%). Also, 84.73% and 86.87% of the patients enrolled under the pre T&T and T&T groups had never smoked cigarette.

Finally, the proportions of participants with disclosed HIV status were similar in pre T&T and the T&T groups (79.94% vs. 78.99%). A summary of the sociodemographic characteristics of the subjects is displayed in Table 1.

Table 1: Socio-demographic characteristics of the subjects.

	Pre-T&T group		T&T group		
Variable	N	%	N N	%	P value
Age in years (n=824)	-,	, •		, •	
19-30	85	20.63	89	21.60	
31-40	147	35.86	146	35.44	-
41-50	115	27.91	108	26.21	0.93
>50	65	15.78	69	16.75	
Gender (n=831)					
Male	140	34.15	147	34.92	0.01
Female	270	65.85	274	65.08	0.81
Marital status (n=775)					
Married	142	37.87	152	38.00	
Unmarried	174	46.40	172	43.00	0.45
Divorced	21	5.60	51	12.75	0.45
Widowed	38	10.13	25	6.25	
Employment status (n=743)					
Employed by an institution	86	24.78	81	20.40	
Self-employed	152	43.80	171	43.07	0.21
Unemployed	109	31.41	145	36.52	
Income (n=589)					
<50,000FCFA	109	73.64	148	79.46	
50-100000fcfa	31	12.02	45	13.06	0.03
>100,000fcfa	37	14.34	23	6.95	
Level of education (n=776)					
None	28	7.61	29	7.11	
Primary	131	35.60	147	36.03	0.00
Secondary	160	43.48	199	48.77	0.09
University	49	13.32	33	8.09	
Time of travel (n=607)					
0-30 minutes	135	51.53	193	55.94	
0.5-1 hour	79	30.15	98	28.14	
1-2 hours	34	12.98	34	9.86	0.56
>2hours	14	5.34	20	5.80	
Alcohol (n=621)					
No	174	36.96	102	38.84	0.63
Yes	134	63.04	211	61.16	
Smoking (n=743)					
No, former smoker					
	34	9.80	38	9.60	
No, never smoked	34 294	9.80 84.73	38	9.60 86.87	0.43

Continued.

Variable	Pre-T&T group		T&T group		Dyolyo
	N	%	N	%	P value
HIV status Disclosure (n=720)					
Yes	69	20.06	79	21.01	0.75
No	275	79.94	297	78.99	0.75

P value for Chi square test.

Table 2: Clinical outcomes.

Variable	Pre T&T	T&T	P value (chi square)
CD4 (median)			
Initiation	194 (83-367)	225 (101-411)	0.03
at 6 months	335 (214-539)	405 (203-624)	0.50
at 12 months	365 (207-557)	451 (236-637)	0.04
at 24 months	384 (226-747)	518 (240-550)	0.89
BP systolic (mean)			
Month 6	123±28	129±24	0.34
Month 12	133±24	129±21	0.54
Month 24	120±22	124±17	0.41
BP diastolic (mean)			
Month 6	77±15	75±16	0.33
Month 12	81±12	82±14	0.86
Month 24	76±12	81±14	0.17
Weight (mean)			
1 month	64±16	66±15	0.16
6 months	67±14	76±49	0.06
12 months	73±64	70±15	0.26
24 months	69±12	72±17	0.14

HIV clinical characteristics at baseline and follow up care

The median CD4 count values (cells/mm³) at enrolment in the pre T&T and T&T groups were respectively 199 and 239, p= 0.01. There was a general increase in the median CD4 count values from baseline in both groups through months 6, 12, and 24 (326 vs.415 p=0.13; 365 vs.505, p=0.01; 448 vs. 518, p=0.62). The values were generally higher in the group enrolled during the test and treat and the difference was significant at initiation (month 1) and at 12th month.

Amongst those who did HIV viral load test, the proportions with undetectable viral load in the pre T&T and the T&T groups at months 6, 12, and 24 were; 59.09% vs. 82.86%, p=0.04; 59.26% vs.72.09%, p=0.18; 76% vs.83.33%, p=0.51. Viral Suppression proportion was significantly higher at month 6 amongst those enrolled under the T&T.

Regarding blood pressure (BP), the mean systolic BP at 6, 12 and 24 months were similar in the pre-T&T and the T&T group (123 ± 27.79 vs. 129 ± 24.26 ; 133 ± 23.64 vs. 129 ± 20.61 ; 120 ± 21.91 vs. 124 ± 17.45). Similar trends were observed for the diastolic BP (77 ± 115.26 vs. 75 ± 15.73 ; 81 ± 12.1 vs. 82 ± 13.73 ; 76 ± 12.33 vs. 81 ± 13.7).

The pre T&T and T&T baseline mean weights were 64 ± 15.64 and 66 ± 14.76 , p=0.16. The means generally increased through 6,12 and 24 months in both groups, though generally higher in the T&T group, with a significant difference at month 6 (67.04 vs.75.87, p=0.00).

A summary of the clinical outcomes is displayed in Table 2 below.

ART uptake and retention

ART uptake proportion was significantly higher in the T&T group compared to the pre T&T group (98.09% vs. 95.39%, p=0.02).

With regards to the respect of clinical appointment dates, patients in the pre T&T group were more likely to respect their appointment dates. The proportions of patients in the pre T&T and T&T groups who respected their clinic dates at months 6, 12, 24 were: 76.62% vs. 62.67%, p=0.00; 67.95% vs. 56% p=0.01; 53.49% vs. 42.98%, p=0.03.

There was a general decline in ART retention over time in both groups. Retention at month 3, 6, 12 and 24 in the pre T&T and T&T groups were respectively: 87.02 vs. 89.10, p=0.35; 84.95 vs. 81.57, p=0.19; 78.24% vs. 77.25%, p=0.73; 85.76 vs. 78.83, p=0.01. However, the

decline in retention was higher in the T&T group, with a significant difference at 24 month. Figure 2 illustrates the

ART retention trends in both groups and ART uptake and retention statistics are summarized in Table 3.

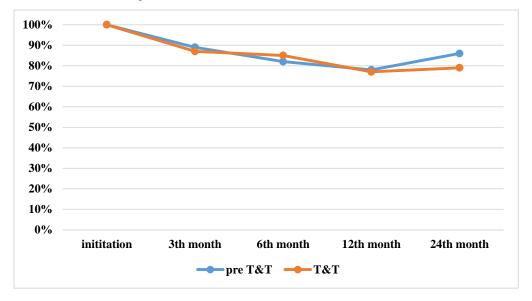


Figure 2: HIV test and treat and pre-test and treat retention trends.

Table 3: ART uptake and retention outcomes.

¥7	pre-T&T	pre-T&T group		coup	p-value	
Variable	N	%	N	%	(Chi Square)	
ART uptake (n=831)						
Yes	393	95.39	411	98.09	0.02	
No	19	4.61	8	1.91	0.02	
Duration diagnosis ART uptake (n=740)						
<2 weeks	171	48.17	215	55.84	0.03	
>2 weeks	184	51.83	170	44.16	0.03	
Respect of clinical follow-up visits						
Month 6 (n=456)						
yes	177	76.62	141	62.67	0.001	
No	54	23.38	84	37.33	0.001	
Month 12 (n=459)						
Yes	159	67.95	126	56.02	0.008	
No	75	32.05	99	44.08	_	
Month 24 (n=442)						
Yes	115	53.49	98	42.98	0.04	
No	99	46.51	130	57.02	0.04	
ART retention at month 3 (n=829)						
Retained	362	87.02	368	89.10	0.25	
Not retained	54	12.98	45	10.90	0.35	
Retention at month 6 (n=819)						
Retained	350	84.95	332	81.57	0.19	
Not retained	62	15.05	75	18.43		
Retention at month 12 (n=809)						
Retained	320	78.24	309	77.25	0.73	
Not retained	89	21.76	91	22.75		
Retention at month 24 (n=786)						
Retained	338	85.79	309	78.83	0.01	
Not retained	56	14.21	83	21.17		

DISCUSSION

The WHO recommended test and treat (T&T) strategy to avert the HIV/AIDS pandemic was implemented in Cameroon since 2016. There is, however, paucity of literature on the ART outcomes of the strategy in Cameroon. This study assessed the ART uptake and retention outcomes of the HIV T&T strategy in three Cameroon HIV clinics. The study found a higher ART uptake proportion and earlier ART initiation amongst patients enrolled under the HIV T&T strategy. However, ART retention at 24th month was lower amongst patients enrolled under the T&T guidelines. This study underscores a need for more extensive assessment of the ART outcomes of T&T strategy in diverse settings in order to contextualize ART programs to favour optimal outcomes.

Baseline comparisons revealed no significant difference in sociodemographic characteristics between patients enrolled in care in the study clinics under the pre T&T and the T&T guidelines. This was expected as the records analysed came from the same general population and within a similar time period (six months period including a three month period before and a three month period following the T&T strategy implementation).

As expected, some differences were observed in the clinical characteristics between the pre-T&T and the T&T group. The higher mean CD4 count at baseline and at 12th month in the T&T group is consistent with expectation as the T&T strategy has been associated with a higher proportion of asymptomatic patients initiating care. This is further illustrated by the observed higher mean weight amongst patients enrolled under the T&T group.

The very high ART uptake observed in both groups is not surprising as these were patients already enrolled in care in the HIV clinics. Patients already linked to care in HIV clinics are more likely to initiate ART. 13,14 The higher ART uptake proportion (98.09 vs. 95.39, P=0.02), and higher proportion of patients who initiate ART within 2 weeks following HIV diagnosis in the T&T group is consistent with other studies, reporting higher ART uptake and earlier ART initiation under the T&T guidelines. 15,16 These constitute very important positive outcomes of the HIV T&T strategy in Cameroon. It was interesting to note that the difference in ART uptake proportion and duration between HIV diagnosis and treatment was not very huge. This could be explained by the progressively less restrictive CD4 based ART eligibility criteria over the years, from <200 through 350 to <500 cells/mm3 just before the implementation of the T&T around mid-2016 in Cameroon.

This study found a general decline in ART retention in the T&T and the pre-T&T patients over a 24 month period, a typical trend in ART programs.¹⁷ Retention in care is expected to decrease over time as some patients die along the line while others get lost to follow up (LTF)

or transferred out. The highest decline in retention was recorded during the first six months in both the T&T and the pre T&T groups, with a the T&T group recording a greater than three times higher decline in ART retention during the 3rd to 6th month (7% vs. 2%), and significantly lower retention at 24th month. The higher decline in ART retention during first 6 months has been reported in ART programs.¹⁸ This disproportionate decline in retention raises major concerns and the trends are somewhat contradiction with those reported in other studies. A higher ART retention was reported at 12 under the T&T guideline (83.0% vs. 76.2%) by study in Malawi and by a study in rural South Africa (82% vs. 75%). 19,20 It should however be noted that the higher T&T retention reported in the two aforementioned studies was at 12 month, while this study reported lower T&T retention at 24 month, and similar ART retention proportions at 12 month. The 12 month retention reported by this study is consistent with the results of ART retention survey in the ten regions of Cameroon conducted in 2015.²¹ A meta-analysis in 2017 found a trend toward an increased risk of being lost to follow-up at 6 months with rapid ART initiation.²² The differences observed in the retention outcomes of the T&T strategy could be related to contextual differences in the populations of persons living with HIV as well as the T&T strategy's implementation approach. Rushing of patients to initiate ART without adequate counselling, overcrowding and long waiting time in HIV clinics were reported to be linked with higher attrition rates observed at the early phase of the T&T implementation in some clinics in Cameroon.²³

It was also noted that the higher proportion of asymptomatic patients enrolled under the T&T guidelines, constitutes a subgroup associated with higher ART attrition. This is seemingly true for this study which found a significantly higher median CD4 value in patients enrolled under the T&T. Lower retention has been reported in patients with higher CD4 count and gaining weight in Nigeria and South Africa. In the same light, "feeling healthy" was reported as a barrier to ART initiation and engagement by a study in Cameroon and another in Mozambique. These suggest that proper counselling, therapeutic education and close follow up amongst healthier patients initiating ART maybe be helpful in improving ART retention, especially under the T&T context.

CONCLUSION

This study found both positive and negative implications of the HIV T&T strategy on ART uptake and retention outcomes. While it was associated with a higher ART uptake and earlier initiation on one hand, lower ART retention was recorded on the other hand. ART attrition during the 3rd to 6th month was more than three times higher amongst patients enrolled under the T&T guidelines. Improved ART uptake proportion and earlier ART initiation constituted positive outcomes of the HIV test and treat strategy. However, the strategy would be

more effective if retention outcomes are also improved. These findings underscore the need to assess patterns and determinants of early attrition under the test and treat guidelines in order to inform strategies to improve ART retention outcomes.

Limitations of the study

This study was a retrospective chart review, as such was subjected to some limitations associated with retrospective chart reviews. An explicit protocol and well defined inclusion and exclusion criteria were used and data abstracted from all the medical charts meeting the study inclusion criteria to minimized risk of selection bias. Standardized piloted abstraction forms were used and the variables to be collected were well defined in order to minimize misclassification bias. The data was abstracted across the three study sites by a trained team. Records with missing data on the outcome variables were not included in the analysis and this did not affect the results of the study as there were very few missing records which were random.

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