

Major Article

Treatment adherence in patients living with HIV/AIDS assisted at a specialized facility in Brazil

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Abstract

Introduction: In the 1990s, Brazil adopted a public policy that allowed for universal, free access to antiretroviral therapy (ART). Since then, treatment adherence has become a new challenge for administrators of sexually transmitted disease/acquired immunodeficiency syndrome (STD/AIDS) policies. This study quantified adherence to ART and verified whether there is an association between sociodemographic variables and clinical/laboratory data in human immunodeficiency virus (HIV)-infected patients. **Methods:** This was a cross-sectional, exploratory study with a quantitative approach that was conducted over 8 months. The target population contained patients who were assisted at the ambulatory care facility specialized in STD/AIDS of a mediumsize city located in Northwest São Paulo. In order to verify the level of adherence to ART, a validated CEAT-VIH (Assessment of Adherence to Antiretroviral Therapy Questionnaire) questionnaire was used. Sociodemographic aspects and clinical/laboratory data were obtained from the medical records. The results were analyzed using the Student's t-test and Pearson's coefficient. Results: Herein, 109 patients were interviewed, 56% of whom were male. The age of the population ranged 18-74 years (mean 45.67 years). Adherence to ART was classified as insufficient in 80.7% of cases. There was an association between ART adherence and presence of symptoms and/or opportunistic infection (p=0.008) and economic status (p<0.001). Conclusions: Adherence to ART among HIV carriers cared for by the public health system is low. Patients who reported a favorable economic status and those without symptoms and/or opportunistic infection demonstrated greater treatment adherence than those who needed to take more than 3 pills a day.

Keywords: HIV Seropositivity. Medication adherence. Unified Health System.

INTRODUCTION

In the 1990s, Brazil adopted a public policy that allowed for universal access to antiretroviral therapy (ART), with free distribution of medication to all people infected with human immunodeficiency virus (HIV). The strategies focused on the prevention of new cases of HIV infection and epidemic control have resulted in a reduction of morbidity, mortality, and hospitalization rates, and thus an increase in life expectancy¹.

As a consequence of this health policy, treatment adherence has become a new challenge for administrators of sexually transmitted disease (STD)/acquired immunodeficiency syndrome (AIDS) policies as well as for those health services directly involved in assisting HIV carriers in Brazil2.

Viral suppression can be achieved as long as there is consistent adherence to ART among the patients. However,

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when medication is taken incorrectly or to an insufficient degree (lapses, not complying with the days and times, etc.), it facilitates virus replication, which may undergo mutations, generating multiresistant strains and consequent therapeutic failure and reducing future treatment options^{3,4}.

In Brazil, although a few studies have been conducted to verify the rate of adherence to antiretroviral therapies, this evaluation is currently one of the most impactful efforts to reduce complications and improve the quality of life of people living with HIV/AIDS. Therefore, the aim of this work was to determine the level of adherence to pharmacotherapeutic treatment and ascertain whether there is an association between sociodemographic variables and the clinical data of HIV carriers cared for by the public health system.

METHODS

This was a cross-sectional, exploratory study with a quantitative approach. The target population was composed of HIV-positive patients registered at an outpatient care facility specializing in STD/AIDS, in a medium-sized city in the northwest region of the Brazilian State of São Paulo. The inclusion criteria were as follows: having an officially registered and confirmed HIV diagnosis, age 18 years or older, and agreeing to take part in the study. The exclusion criteria were illiterate status, being in a situation of deprivation of liberty (arrested), and cognitive difficulty. Subjects were recruited via convenience sampling as they came to the facility for treatment. The data collection period was 8 months. Data were collected when patients visited the facility for their medical appointments, in a private room, before the appointment began.

The CEAT-VIH (Assessment of Adherence to Antiretroviral Therapy Questionnaire), validated in Brazil by Remor et al.⁵, was used to determine adherence to treatment. The CEAT-VIH is a self-reported questionnaire comprising 17 questions that assess the level of adherence to antiretroviral treatment. It has a multidimensional character, as it comprises the principal factors that may shape treatment adherence behavior, and consists of five domains: treatment observance (following the instructions to take the medication correctly), history of treatment non-adherence, doctor-patient communication, personal beliefs (treatment expectations), and treatment satisfaction. The final score is obtained by summing all items, the minimum possible score being 17 and maximum 85. According to the CEAT-VIH, if the gross score is ≥80, the patient is considered highly adherent, and if <80, non-adherent (Table 1).

Sociodemographic characteristics and clinical and laboratory data were obtained by analyzing the medical records. Examination results used in this study were those closest in date to the participation in the interviews. Sociodemographic variables included age, sex, level of education, marital status, occupation, ethnicity, sexual orientation, and perception of economic status. The variables related to HIV infection included viral load, CD4 count, presence of symptoms or opportunistic infection in the past few months, number of pills taken, and mode of infection.

The assessment of sociodemographic and (HIV-related) clinical/laboratory variables regarding adherence to treatment was performed using the Student's t-test or the analysis of variance, in the event that there were two or three categories, respectively. Pearson's coefficient was used to verify the correlation between the numeric variables and treatment adherence scores. Statistical tests were performed with a level of significance of 5%.

Ethical considerations

For the analysis of the data, the software programs Microsoft Office Excel 2016 and Statistica (StatSoft Inc.), version 10, 2011 were employed. The study was approved by the Human Research Ethics Committee (Process FOA-02411/2011) and written informed consent was obtained from each participant. All procedures followed were in accordance with the ethical standards of the responsible committee on human experimentation (institutional and national) and with the Helsinki Declaration of 1975, as revised in 2000.

RESULTS

A total of 109 patients were interviewed, 56.0% of whom were male. The subjects' ages ranged from 18 to 74, with the mean age being 45.7. Adherence to antiretroviral treatment was classified as insufficient in 80.7% of cases (**Table 2**).

TABLE 1
Classification of level of adherence to antiretroviral therapy.

Classification of adherence	Adherence score
High adherence	≥80
Non-adherence (insufficient)	<80

TABLE 2

Classification of level of adherence to treatment among HIV carriers according to the CEAT-VIH test.

Adherence score	Classification	Number	Percentage
≥80	High adherence	21	19.3
<80	Non-adherence	88	80.7
Total		109	100.0

HIV: human immunodeficiency virus (HIV); CEAT-VIH: Assessing Adherence to Antiretroviral Therapy.

Table 3 shows a selection of measurements depicting total adherence scores according to the categories established for the variables studied. Owing to the small number of individuals in some initially planned categories, we chose to group the subjects into more relevant groups using only two or three categories. The table also displays the p-values obtained from the application of the analysis of variance t-test or F-test, where the variable has 2 or 3 categories, respectively. Our results showed that individuals who reported the absence of opportunistic infections had higher treatment adherence (p=0.008), as did individuals with a better economic status (p<0.001). Moreover, patients who needed to take more than three pills a day did not demonstrate adverse treatment adherence (p=0.053), although the result is marginal.

Pearson's coefficient was used to verify correlations between the variables (age, time elapsed since diagnosis, CD4 count, and number of pills), global adherence, and the five dimensions that made up the questionnaire. The data showed a negative correlation for time elapsed since diagnosis and number of pills, and a positive correlation with regard to the CD4 count (**Table 4**).

DISCUSSION

In our study, the factors associated with adherence to ART were non-existence of opportunistic infection, economic status, and the number of pills taken daily. Considering the findings of this study, we ascertained that the patients' sociodemographic profiles (age and sex) resemble that of patients from other studies⁶⁻⁸ with similar goals and which used the same tool (CEAT-VIH) to investigate drug therapy adherence in different regions of the country (northeast and south).

The age range of people who live with HIV/AIDS (PLHA) in Brazil differs from that in scientific research data as well as data provided by government information systems. The mean age obtained in the present study was 45.67 years, in agreement

TABLE 3

Frequency and percentage of individuals in the two categories of variables in this study and descriptive statistics of treatment adherence scores.

Categories	n (%)	M (Sd)	Md (1Q; 2Q)		
Age (years)	II (/0)	IVI (DU)	1114 (14, 24)	p	
18 to 39	35 (32.1)	72 (9)	73 (66; 79)	0.127	
≥40	74 (67.9)	74 (7)	75 (71; 79)	0.12/	
Sex Sex	74 (07.3)	74(7)	73 (71, 79)		
females	48 (44.0)	72 (8)	73 (68; 78)	0.100	
males	61 (56.0)	72 (8) 74 (8)	75 (72; 79)	0.100	
Level of education	01 (30.0)	74 (8)	13 (12, 19)		
middle school	49 (44.9)	73 (7)	74 (70; 77)	0.756	
high school	44 (40.4)	74 (7)	75 (70; 79)	0.750	
college degree	16 (14.7)	74 (7)	80 (71; 81)		
Marital status	10 (14.7)	74 (13)	80 (71, 81)		
	40 (26.7)	72 (10)	75 (72: 70)	0.681	
single married	40 (36.7) 39 (35.8)	73 (10) 74 (7)	75 (72; 79) 75 (71; 79)	0.081	
divorced/widowed					
	30 (27.5)	72 (6)	73 (68; 78)		
Occupation employed	40 (44 0)	75 (9)	76 (72, 70)	0.071	
• •	48 (44.0)	75 (8)	76 (73; 79)	0.071	
unemployed Economic status (Self perception)	61 (56.0)	72 (8)	73 (68; 78)		
` • • ′	21 (10.2)	(((0)	(0 ((2, 72)	<0.001	
worse than the others	21 (19.3)	66 (9)	69 (63; 73)	<0.001	
equal or better	88 (80.7)	75 (7)	76 (72; 79)		
Ethnicity	(4 (50.7)	74 (0)	7((72, 70)	0.205	
white	64 (58.7)	74 (9)	76 (72; 79)	0.385	
brown/black	45 (41.3)	73 (7)	73 (68; 79)		
Sexual orientation	00 (02 ()	72 (0)	75 (70, 70)	0.252	
heterosexual	90 (82.6)	73 (8)	75 (70; 79)	0.353	
homo/bisexual	19 (17.4)	75 (10)	75 (72; 81)		
Mode of transmission	04 (74.4)	(0)	-4/50 -00		
sexual	81 (74.3)	73 (8)	74 (69; 79)	0.463	
blood	12 (11.0)	75 (8)	78 (73; 80)		
unknown	16 (14.7)	76 (4)	76 (73; 79)		
Opportunistic infection			,		
no	85 (77.9)	74 (7)	75 (72; 79)	0.008	
yes	24 (22.1)	70 (9)	71 (64; 76)		
CD4 (cells/mm³)					
≤350	22 (20.2)	72 (10)	76 (69; 78)	0.409	
>350	87 (79.8)	74 (7)	74 (70; 79)		
Viral load (copies/mL)					
≤50	75 (68.8)	74 (7)	75 (71; 79)	0.092	
>50	34 (31.2)	71 (10)	75 (68; 78)		
Pills (n°)					
≤3	63 (57.8)	75 (7)	75 (72; 79)	0.053	
>3	46 (42.2)	72 (9)	74 (68; 77)		

N: frequency; %: percentage; M: mean; Sd: standard deviation, Md: median; Q: quartile; CD4: cluster of differentiation 4; ANOVA: analysis of variance. *Significant at the level of 5% (Student's t-test was applied for dichotomous variables or ANOVA in the case of two or more categories)

TABLE 4					
Descriptive statistics of numeric variables and their correlation coefficients with global adherence and other domains.					

Variable	M (Sd)	Md (1Q, 2Q)	D1	D2	D3	D4	D5	D6
Age (years)	45 (12)	47 (36; 53)	0.065	0.073	0.129	-0.062	0.077	-0.069
Time (years)	9 (7)	8 (3; 13)	-0.132	-0.134	-0.005	-0.058	-0.242*	0.031
CD4 count (cells/mm³)	646 (335)	635 (426; 869)	-0.037	-0.051	-0.138	0.251*	-0.064	-0.013
Pills (n°)	3 (2)	3 (3; 4)	-0.213*	-0.220*	-0.173	-0.014	-0.229*	-0.019

M: mean; Sd: standard deviation, Md: median; Q: quartile; D1: global adherence; D2: treatment observance; D3: antecedents of non-adherence; D4: doctor-patient communication; D5: personal beliefs; D6: treatment satisfaction; CD4: cluster of differentiation 4. *Significant correlation at the level of 5%.

with other national studies⁸⁻¹⁰, in which the largest proportions of PLHA were aged between 30 and 49 years. However, data presented in the last HIV/AIDS epidemiological bulletin, published by the Ministry of Health in 2016, show that the highest concentration of HIV cases in Brazil is in individuals aged between 25 and 39 years, for both sexes¹¹. In the last few decades, there has also been an increase in the number of children between the ages of 0 and 14 infected by HIV as a result of vertical transmission, due to the increase in infected women of childbearing age (15 to 49 years)¹².

The low prevalence of HIV infection in the male sex demonstrates that the disease profile has been changing, in which feminization is evident¹³. Although at the beginning of the epidemic homosexual men were most commonly affected by HIV, the current proportion of infected men and women is very similar^{14,15}. The same is true for the majority of the participants declared as heterosexual (82.6%) and not having a stable partner-single (36.7%) (**Table 3**).

This new reality demonstrates the transitional nature of the disease, in which the idea of there being specific risk groups (homosexuals and users of injectable drugs) has given way to the notion of risky behavior (unprotected sex)16. This fact can also be demonstrated in the results of this investigation, as the majority of participants (74.3%) reported having been sexually infected (Table 3). This mode of infection is responsible for most new cases of infection worldwide. The frequency of heterosexual transmission is growing and women are more becoming vulnerable to being infected by their male partners rather than the other way around¹⁷. For many couples, unprotected sex can be interpreted as intimacy and confidence, resulting in a barrier to the use of condoms^{17,18}. Therefore, measures to raise awareness among the general population to prevent the risk of new infections must be predicated principally on risky sexual behavior.

As for the level of instruction, 55.1% of the participants completed secondary education or higher (graduate-level and/ or postgraduate education), and the highest percentage of Brazilians infected by HIV in 2016 had completed secondary education¹¹. This fact contradicts the country's tendency towards the pauperization of the epidemic, i.e. the growing dissemination of HIV in the population with low levels of education⁶. Thus, it is important that the campaigns for awareness and prevention in

combatting HIV be directed to the whole population, regardless of social vulnerability.

One of the most important findings of this study was the high level of non-adherence to antiretroviral treatment (80.7%). Other similar studies, conducted in different regions of Brazil, which used the same tool to observe adherence to ART, found lower values, such as that of Moraes et al.8, who observed an *unsatisfying* and *low* adherence rate in 71.3% of patients investigated, while Galvão et al.19, found inadequate adherence in 51.3% of individuals. Jacques et al.6, revealed, in their study, a high level of adherence (75.7%). Several factors may significantly affect adherence to treatment such as patient traits (sociodemographic and psychosocial factors), patient-health professional relationships (confidence and confidentiality), health-related factors, disease-related characteristics (HIV-related symptoms), and therapy-related factors (number of pills and side effects of medication)²⁰.

Those who presented no symptoms and/or opportunistic infection demonstrated better adherence. Therefore, patients who correctly take their medication present better systemic conditions, as would be expected, because the efficiency of ART depends directly on patient adherence. In order to maintain an undetectable viral load, the patient must take at least 95% of the prescribed medication⁶, which leads to a decrease in hospitalization rates due to opportunistic infection, as well as a decrease in mortality rate²¹.

We also observed an association between economic status and treatment adherence: the better the economic status, the higher the adherence. Probst et al. ²² performed a meta-analysis in order to ascertain the influence of socioeconomic status on the HIV mortality rate in South Africa and observed that lowincome individuals had a 50% higher chance of dying from HIV infection than people with a better economic status. Thus, people receiving a low income constantly deal with limited resources, which may negatively interfere with self-care of the disease. AIDS treatment is complex and involves several variables such as medical follow-up, withdrawal of medication at the health facility, regular use of the medication at the times prescribed by the doctor, and collection of blood for supplementary examinations, among other forms of care, according to the patient's health. All these factors may negatively interfere with the working routine of seropositive patients and, as a result, self-care may be neglected.

Moreover, there was a negative correlation between the number of pills to be taken daily and global adherence to treatment; individuals who needed to take more than 3 ART pills a day showed lower treatment adherence. The use of multiple medications exposes the patient to a more complex treatment, which requires greater attention, memory, and organization with regard to the times to administer the drugs²³. Moreover, the possibility of triggering side effects increases²⁴. For each pill taken, the risk of non-adherence increases by 12%²⁵.

In addition, there was a negative correlation between time elapsed since diagnosis and personal beliefs. Patients who were diagnosed with HIV more recently presented positive expectations of the treatment. This may be explained by the current advances in medicine in combatting HIV, by the free distribution of medication by the public health system, and by the ease of access to treatment. All these factors converge towards a decrease in mortality rates and hospitalization due to opportunistic infections¹. At the beginning of the epidemic, the image associated with the seropositive patient was that of fragile people in a terminal state in hospital beds, waiting to die. Nowadays, owing to the chronification of HIV infection, this association is no longer applicable²⁶.

The health professional has an important role, together with the patient, in terms of maintaining and adhering to ART. The results obtained in this study demonstrated a positive correlation between the number of CD4 lymphocytes and better professional-patient communication. This leads us to infer that patients with better systemic conditions are more satisfied with the treatment received from the professional team that is caring for them. Professional-patient relations are a facilitating agent in adherence to therapy because, as the patient's doubts and needs are addressed and supported, a trusting relationship and a feeling of belonging are created, which is fundamental to the success of such a complex therapy²⁷. Nevertheless, it is important to emphasize that the patient's level of adherence to ART cannot be definitive in nature as it may vary at any time during the treatment. Therefore, adherence must be constantly stimulated by the health professionals, who must always be aware, in order to take co-responsibility for either the success or the failure of therapy⁶.

For PLHA in Brazil, the national health policy provides universal access to treatment through the free distribution of medication, multidisciplinary care at specialized facilities, as well as prevention campaigns and rapid diagnosis tests. Nonetheless, if adherence to treatment is not high enough for ART to be effective, all this investment may be jeopardized since the irregular use or the insufficient number of doses may lead to the development of resistant HIV viruses^{28,29}. Thus, we must highlight the importance of this study, as the constant monitoring of adherence to ART in different regions of the country can direct the government's actions so that the investment is not jeopardized.

The data obtained show that adherence to ART is low in those patients who attend the outpatient care facility (SAE/SUS) investigated in this study. The patients who reported having a favorable economic status and those with no symptoms or

opportunistic infection presented better adherence to ART, while the adherence of those who needed to take more than 3 pills a day was worse. These results indicate the need to intervene with public actions directed towards the population in question, seeking to contribute with increased adherence to drug therapy. Even though the patient is the focus of the treatment, the team of health professionals must take co-responsibility for this process.

Some limiting factors should be considered, such as the fact that the results of this study cannot be generalized to other locations; however, it is important to stress the need to compare the results of this study with those of other studies with similar goals conducted in Brazil and other countries.

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Conflict of interest

The authors declare that there is no conflict of interest.

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