Research Article

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Antiretroviral treatment adherence and associated factors among people living with HIV in developing country, Myanmar

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ABSTRACT

Background: The provision of antiretroviral therapy (ART) has been started in Myanmar, since 2005 by both public and private sectors. Adherence is one of the major critical issues in the clinical management of HIV-infected patients receiving ART. This study aims to determine the level of adherence to ART and possible associated factors in public ART centers, Yangon Region, Myanmar.

Methods: MT his cross-sectional analytical study was conducted between May to July 2015. A total of 425 people living with HIV (PLHIV) were interviewed with pretested structured questionnaires. The multi-method tools were used to access the level of ART adherence. The generalized estimating equation (GEE) model and multiple logistic regression models were administered to determine the possible associated factors to ART adherence.

Results: Only 76.24% (95% CI=72.17-80.29) of the respondents had optimal ART adherence (≥95%). The respondents of aged more than 40 years were unlikely to adhere ART when compared with younger age (AOR=0.01, 95% CI=0.003-0.05). Those patients who had not satisfied with ART provider services (AOR=0.04, 95% CI=0.02-0.12) were also significantly less likely adhering to ART. However, non-smokers and being taking regular exercise were found to be more likely to adhere to ART.

Conclusion: About a quarter of the PLHIV had suboptimal ART adherence. The older age group and poor patient-provider relationship were found to be the barrier to ART adherence.

Keywords: Treatment for HIV/AIDS, Antiretroviral therapy adherence, Myanmar

INTRODUCTION

Acquired immunodeficiency syndrome (AIDS) is one of the best known chronic diseases that needs daily medications provided by healthcare system of different regions of the world in order to control the associated complication and maintain good health. Although the clinical outcomes of HIV infected patients have been improved by ART, maintaining an optimal adherence to ART is a great challenge. Moreover, adverse effects of ART, complexity of the dosing and socio-economic factors also contribute the challenges to the patients and often influence the ART adherence. Optimal adherence to ART suppresses the viral replication and less

likelihood of transmission of HIV to uninfected persons. Furthermore it can reduce morbidity, mortality, also health care costs for PLHIV and also related to the quality of life of those patients. However, almost one third of patients has been missing their doses for ART every 5 days and ART adherence remains a great challenge for many persons receiving ART, and thus the benefits and outcomes of ART are not fully accomplished. 8-10

Treatment adherence is the quality of sticking to particular instructions by the health care provider including to follow dietary instructions, clinic appointments and lifestyle adjustment. That is why

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adhere to ART is a very complex and dynamic phenomenon which involves the patients, health care providers, the health system within which they operate. Adherence rate of up to 95 percent is necessary for HIV viral suppression and with 80 percent of adherence can maintain the viral load undetectable over six months. Therefore, accurate and reliable measures of ART adherence and better understanding of the influencing factors on adherence are essential for clinicians to identify patients who need assistance for taking pills, to develop and evaluate effective interventions to enhance adherence, and to evaluate clinical outcomes and making treatment decisions. ¹³

In Myanmar, AIDS is one of the diseases of National concern. The ART provision has started since 2005 and has covered 139 ART Centers/sites including government hospitals and AIDS/STD clinics in public sectors. The number of PLHIVs is estimated about 198,811 for 2013. About 75,000 AIDS patients have been treated for ART in 2014 through coordinated efforts of fifteen implementing partners. According to the fourth edition of National guidelines for the clinical management of HIV infection, with the eligible criteria of CD_4 count < 500, the need for ART in Myanmar is estimated to be more than 120,000 in 2014.

Myanmar has planned to scale up for over 100,000 patients in 2015 and over 111,000 patients by the end of 2016. In order to achieve the ambitious scale up plan, the National AIDS Program has started countrywide ART decentralization since 2013 and has a plan to cover over 100 townships by the end of 2015, with the aim of expanding area coverage, decongestion of ART center work load and improve accessibility for community by reducing the travel time and costs. Although a few studies had been undertaken on ART adherence in different sites of Myanmar, published literature focusing on adherence to ART and its determinants were limited. 15-16

This study aims to determine the level of adherence to ART and possible associated factors to provide the baseline data for policy makers in ART provision and adherence counseling services and decentralization of ART plan in Myanmar.

METHOD

This cross-sectional analytical study was conducted at all public ART centers of Yangon Region. The study populations were PLHIVs attending the outpatient department of the ART centers and receiving ART with first line drugs between May and July 2015. Those patients who gave informed consent, aged 18 years and above and have been taking ART for at least 3 months were eligible for this study. The PLHIVs attending at the prison ART center and those patients who cannot communicate properly because of ill health were excluded in this study.

The sample size was estimated based on the multiple logistic regressions formula¹⁷ and was calculated by PASS software version 13. Although the calculated sample size was 204, the calculated sample size was multiplied by inflation factor to avoid the clustering effect. Therefore the final samples to be collected were 425. Yangon Region was selected as the study area because of being the largest ART center and HIV/AIDS hospitals are situated in Yangon. There were altogether 9 public ART centers in Yangon Region. Apart from prison ART center, 5 percent of the total attendees from each center were recruited to get the required sample size. The data was collected on the OPD days of each ART center. The eligible samples were selected by applying systematic random sampling from the OPD register book on the day of data collection.

Assessment of ART adherence was carried out by using multi-method tools validated for assessing the ART medication adherence of adult HIV infected patients in resource-constrained setting 18 which consists of self report, visual analogue scale (VAS), pill identification test (PIT) and pill count method. The participants had to answer four questions in self-report form. The questions include patient's recall for any missed dose in the past four days, whether the patient had difficulty to remember to take the medication, any drug holidays and stopping the drug because of side effects. The answer was Yes or No. The answer of No to all questions was scored as high adherence and yes to one question and two or more questions were scored as moderate and low adherence respectively.

Regarding visual analogue scale, the participants had to recall back over the past four days and identified the number of times if he or she either missed a dose or took it at the wrong time. A line with 10 marks from 1 to 10 was drawn and the participants had to mark with a cross on it according to their medication doses that they had taken in the past 4 days. If he or she had taken all medicine doses to point to 10 and if the participants missed all the doses, he or she would point to 0. The percentage adherence would be 40 percent if the scale was marked at 4. The results of VAS were scored as high adherence (95% and more), moderate adherence (75% to 94%) and low adherence (<75%).

In pill identification test, each patient was asked to show his or her issued medication and checked each container and its contents. He or she had to tell the name of the medication, number of pills have been taken per dose, the time for medication have been taken, and any additional instructions. The scores for (PIT) were high adherence (patients who knew the dose, time for medication and additional instructions), moderate adherence (those who knew only dose and time for medication) and low adherence (those who knew only dose or confused).

Adherence from pill count method was calculated by dividing the difference between the dispensed pills and

returned pills by the number of pills that have to be taken during the same period. Each patient was asked to recall about the dispensed pills in previous month and returned pills or extra pills when he or she came back for follow up. If leftover medication could not recall, then the calculation was invalid. In this multi-method adherence tool, the overall resultant adherence level was categorized into high level (95% or more), moderate level (75% to 94%) and low level (less than 75%). The optimal adherence in this study was considered as adherence of 95% or more and suboptimal adherence as lower than 95%.

In this study, the dependent variable is ART adherence, which is categorized into optimal and suboptimal. Age, sex, education, occupation, marital status, ART regimen, duration of ART, CD4 count co-morbidities, patient provider relationship, disclosure status, exercise, alcohol drinking and smoking were independent variables.

The data were collected by face-to-face interview with pretested structured questionnaires except self-report form for assessment of ART adherence. Prior to the interview, the questionnaires were translated into Myanmar and back translated into English by an independent translator. One day training was given to four research assistants for data collection about the purpose and facts to follow while asking the questionnaires. All interviews were performed in the separate rooms of the ART centers to ensure privacy and confidentiality. The data were collected after informing about the purpose of the study, risk, benefits and assuring confidentiality to those who were eligible for this study. Immediately after interview, the error, consistency and completeness of the questionnaires were checked to assure the quality of the data. Reviewing of medical records was also done to record the latest CD4 counts and associated co-morbidities.

The data were entered with Epi-data (version 3.1) and analyzed by using the Stata program version 13.0 (Stata Corp, College Station, TX). ART adherence of the participants with 95% CI was calculated. To determine the factors associated with ART adherence, odds ratios (ORs) and 95% CI were estimated using multiple logistic regression model. Significant factors in a bivariate analysis with p < 0.25 were included in a multiple logistic regression analysis. Since the data collection was conducted at the ART centers; cluster level analysis was performed to adjust for similarities within each center, using the generalized estimating equation (GEE) model. All test statistics were two-sided and a p-value of less than 0.05 was considered as statistical significant.

Ethical consideration

The Ethical Committee of Khon Kaen University, Thailand had approved this study protocol. Informed consent had taken from all participants prior to interview and all information provided by the participants was confidential and would not be shared to anyone apart from the research team.

RESULTS

Table 1: Socio-demographic characteristics of the respondents.

Characteristics Total (n = 425) Number Percent Gender Male 236 55.53 Female 189 44.47 Age (years) 44.47 44.47	
Gender 236 55.53 Female 189 44.47	
Male 236 55.53 Female 189 44.47	
Female 189 44.47	
rige (Jears)	
< 40 203 47.76	
>40 222 52.24	
Mean (SD) 41.57 (9.62)
Marital status	(>10=)
Single 81 19.06	
Married 234 55.06	
Separate/divorce/widow 110 25.88	
Educational attainment	
Illiterate 6 1.41	
Can read and write 8 1.88	
Primary school 74 17.41	
Middle school 127 29.88	
High school 143 33.65	
University Graduate 67 15.76	
Occupation	
Government staff 29 6.82	
Private employee 62 40.59	
Dependent 86 20.24	
Manual labour 29 6.82	
Own business 197 46.35	
Driver 22 5.18	
Per capita income (MMK)*	
< 10,000 358 84.24	
> 10,000 67 15.76	
Mean (SD) 72891.26 (58141.07)	
Smoking	
Yes 81 19	.06
No 344 80	.94
Alcohol consumption	
Yes 32 7.5	53
No 393 92	.47
Taking regular exercise	
	.88
No 162 38	.12
Disclosure status	
Disclose 411 96	.71
Not disclose 14 3.2	29

(MMK)*: Myanmar currency (Kyats)

Table 2: Medical history and ART related characteristics.

Duration of ART (months) ≤6	Number 141 284	Percentage 33.18
<u>≤</u> 6 > 6		33.18
->6		33.18
	284	
Mean (SD)		66.82
		35.09 (36.59)
Level of knowledge		
<u> </u>	353	83.06
	72	16.94
Missed ART dose in last month		
	26	6.12
	399	93.88
Reasons for missed ART (n=26)		
	13	50.00%
	15	57.69%
	1	3.85%
	1	3.85%
Missed follow up appointment		
	85	20.00
No .	340	80.00
Side effect of drugs		
Yes	116	27.29
No :	309	72.71
Comorbidities		
	75	17.65
	350	82.35
CD ₄ count (mm ³)		
	186	43.76
≥350	239	56.24
Mean (SD)		415.88(251.36)
Travelling time to ART center (minutes)		
	130	30.59
	220	51.76
	75	17.65
Mean (SD)		123.44(132.21)
Waiting time at ART center		
	28	6.59
	247	58.12
	150	35.29
Mean (SD)		178.91 (91.24)
Satisfaction on ART provider service		
	358	84.24
	67	15.76
Relationship with health care providers		
	259	60.94
Not good	166	39.06

A total of 425 PLHIVs from 8 public ART centers were recruited for this study. Majority of them were males (55.53%) and the mean age was 41.57 with the standard deviation (SD) of 9.62. Married group (55.06%) constituted more than divorce and widow group (25.88%) and single (19.6%). More than one fourth of the respondents had attained the high school level and nearly half of them (46.35%) were self-employed. Nearly one

fifth (19.06%) of the respondents reported as current smokers and very few (7.53%) of them were current alcohol drinkers. More than half (61.88%) were reported that they took the physical exercise regularly. Only 14 respondents (3.29%) found to be not disclosed their HIV status (Table 1).

Regarding medical history and ART related characteristics, majority of the patients (66.82%) have been taking ART for more than 6 months duration. Most of them (83.06%) had high level of ART related knowledge score. Only 26 patients (6.12%) had missed their dose of ART one month prior to data collection. The reasons for missed doses were away from home (50%), simply forgot (57.69%), many pills to be taken (3.85%) and medication with social stigma (3.85%).

Twenty percent of the participants had missed their follow up appointment at least one time within 3 months prior to data collection. About one forth (27.29%) of the respondents mentioned experiencing the side effects of

ART and only (17.65%) of them had comorbidities. More than half (56.24%) had the CD_4 count of more than 350 mm³ and the mean CD_4 count was 415.88 with SD (251.36). Majority (84.24%) of the respondents reported that they satisfied with the ART providing services and 60.94% mentioned that they had good relationship with the health care providers (Table 2).

The level of adherence was assessed by using multimethod tools. Nearly one fourth of the patients (21.65%) self-reported that they were difficult to remember to take ART for 4 days recall. Regarding to VAS scores, the ART adherence of more than or equal to 95% was found to be (98.35%).

Table 3: Factors associated with ART adherence using simple logistic regression.

Variable	Total number	Optimal adherence to ART (%)	Crude OR	95% CI	P-value			
Age (years)	1 otal lialisel		Ordae Ore	76 70 01	1 varae			
< 40	203	61.73	1		-0.001			
>40	222	38.27	0.02	0.01 to 0.06	< 0.001			
Gender				0102 00 0100				
Male	236	52.78	1		0.04			
Female	189	147.22	1.62	1.02 to 2.56	0.04			
Per capita income (MM	K)							
≤10,000	358	81.79	1		0.01			
>10,000	67	18.21	2.59	1.19 to 5.62	0.01			
Duration of ART								
≤6 months	141	36.42	1		0.01			
>6 months	284	63.58	0.51	0.31 to 0.86	0.01			
Side effect of drugs								
No	116	75.31	1		0.05			
Yes	309	24.69	0.62	0.38 to 1.00	- 0.05			
Comorbidities								
Yes	75	16.36	1		0.22			
No	350	83.64	1.42	0.82 to 2.48	0.22			
Travelling time to ART	center							
<60 minutes	130	32.10	1					
60 to 180 minutes	220	49.38	0.67	0.39 to 1.12	0.21			
>180 minutes	75	18.52	1.00	0.49 to 2.04	0.21			
Satisfaction on ART pro	ovider service							
Yes	358	95.37	1		<0.001			
No	67	4.63	0.05	0.02 to 0.09	<0.001			
Relationship with health	icare providers							
Good	259	63.27	1		0.08			
Bad	166	36.73	0.67	0.42 to 1.05	0.00			
Smoking								
Yes	81	12.35	1		<0.001			
No	344	87.65	4.85	2.89 to 8.14	\0.001			
Alcohol consumption								
Yes	32	5.86	1		0.03			
No	393	94.14	2.37	1.13 to 4.99				
Taking exercise								
No	263	25.62	1		<0.001			
Yes	162	74.38	10.43	6.11to 17.79				

Table 4: Adjusted odds ratio (AOR) of the factors that were associated with ART adherence and their 95% CI.

Variable	Total number	Optimal adherence to ART (%)	Crude OR	Adjusted OR	95% CI	P-value
Age (years)						
<u>≤</u> 40	203	61.7	1	1	0.002 to 0.05	< 0.001
>40	222	38.27	0.02	0.01	0.003 to 0.03	
Side effect of drugs						
No	116	75.31	1	1	0.23 to 1.16	0.11
Yes	309	24.69	0.62	0.52		0.11
Satisfaction on ART	Satisfaction on ART provider service					
Yes	358	95.37	1	1	0.02 to 0.12	< 0.001
No	67	4.63	0.05	0.04		
Smoking						
Yes	81	12.35	1	1	1.92 to	< 0.001
No	344	87.65	4.85	4.26	9.44	<0.001
Taking exercise						
No	263	25.62	1	1	4.87 to 22.04	<0.001
Yes	162	74.38	10.43	10.36		

Three hundred and twenty four participants (76.24%) could identify the name of the medication, the correct number of pills had been taken per dose, the time for medication had been taken and additional instruction concerning food and water intake. According to the overall resultant adherence level, those with optimal adherence (95% or more) were (76.24%, 95% CI=72.17-80.29, n=324).

Multivariate analysis after adjusting the clustering effect with GEE found that the respondents of aged more than 40 years were unlikely to adhere ART when compared with younger age (AOR=0.01, 95% CI=0.003-0.05). Those patients who had not satisfied with ART provider services (AOR=0.04, 95% CI=0.02-0.12) were also significantly less likely adhering to ART. Non-smokers (AOR=4.26, 95% CI=1.92-9.44) and being taking regular exercise (AOR=10.36, 95% CI=4.87-22.04) were significantly more likely to adhere to ART (Table 4).

DISCUSSION

Adherence usually refers to dose adherence in most studies. However adherence in this study was assessed not only by dose adherence but also scheduling and dietary instructions and then over all resultant adherence was taken into account to consider the level of adherence. The overall adherence level in this study was 76.24%, which was a bit disparity with previous studies 75% 15 and 85% 16 conducted in Yangon Region. In fact the level of adherence in this study was rather higher than other studies across the South East Asia Region where ART adherence varied widely from 57% to 75% 19-21 and the study undertaken in developed country like Latin America, reported the ART adherence as 55%. 22 This may be due to the differences in timing of the study, different measuring tools to assess the adherence level, different settings, and nature of availability of the drugs. Hence comparing the adherences to ART across the

studies had to be done with caution. The reasons for missing their medication doses in current study were away from home, forgetfulness, pills overload and medication with social stigma, which were consistent with previous literature. 23 Because of the self-stigma, the patients dared not taking ART in the presence of others leading to missed doses. A number of studies have been examined for the factors that associated with ART adherence among PLHIV receiving antiretroviral therapy, while some focused on socio-demographic characteristics such as age, gender, education and income, others concentrated on health behavior and cultural beliefs of patients, some emphasized on the patient-provider relationship. The respondents of more than 40 years of age were significantly less likely to adhere ART in this study although several studies reported that there is no consistent correlation between demographic characteristics and level of adherence. ^{24,25} Many potentially debilitating adverse effects of ART have also been shown to contribute to irregular taking of the drugs and deliberately stopping of medication by some patients.²⁶ The results of current study revealed that the side effect of ART was not associated with ART adherence. This could be the reporting of the respondents being experiencing side effects of ART from both optimal and sub optimal adherence groups. In this study, lack of satisfaction on ART provider services significantly influenced the optimal adherence. Patient's satisfaction is achieved by good patient-provider relationship and more confidence in the provider resulting good adherence. Being non-smokers and those who took regular exercise were strong relationship to optimal adherence. These findings were consistent with the study conducted in Lao PDR.²⁷ This may be due to the attitude and adopted life style of the patients.

There were a number of limitations in our study because of its cross-sectional design and assessed the ART adherence at a single time point. In fact, it was impossible to evaluate variations in adherence over time and the extent of generalizability is limited only to those similar patients. Because of cross-sectional nature of the study, it did not allow the cause and effect relationships between various factors and adherence to be addressed. Despite the limitation, this finding was useful for program managers as a base line to develop appropriate strategies to address the issue of adherence from both the clinician and the patient side.

CONCLUSION

In conclusion, the level of adherence in present study was relatively higher than other studies conducting in South East Asia. The older age group and lack of patient's satisfaction were found to be the barriers to ART adherence. Being non-smoker and taking regular exercise were significantly associated with optimal adherence.

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Institutional Ethics Committee

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