

## Research Article

### Factors Associated with Non-Adherence to Antiretroviral Treatment in Adults at Keetmanshoop District State Hospital in Namibia

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#### Abstract

**Aim:** To identify the factors associated with non-adherence to Antiretroviral Treatment (ART) amongst adult patients at a hospital in Namibia.

**Methods:** Quantitative and qualitative approaches were employed, study sample of 112 adults aged 21 years and older at Keetmanshoop District State Hospital participated. Adherence to ART was assessed using the pill count and self-report methods. A questionnaire with a Cronbach value of 0.8 was used. Census and multi-stage sampling was applied. Data was collected from August to September 2016 through structured interviews and patients records review. Correlation of variables was done to remove multicollinearity. Stata version 12 was used for logistic regression to do univariate and multivariate analysis to determine factors associated with missing five or more doses.

**Results:** In the univariate analysis the following factors were associated with missing five or more doses: Viral load OR: 5.6, 95%CI: 2.0-16.0, Employment OR: 3.3, 95%CI: 1.5-7.2, Taking active substance OR: 2.5, 95%CI: 1.3-4.7 and Emotional status OR: 1.6, 95%CI: 1.02-2.6. The ART stocks seemed adequate as the OR for out of stock did not lead to missed doses (OR: 0.6, 95%CI: 0.4-0.9). Factors such as gender, education, religion and marital status were not associ-

ated with missing five or more doses. After multivariate logistic regression the following factors were associated with missing five or more doses; Taking substance adjusted OR: 3.3, 95%CI: 1.4-7.6; WHO clinical stages 2 and 3 adjusted ORs and 95%CI: 4.1 (1.2-13.8) and 6.1(1.5-25.1) respectively and viral load adjusted OR: 6.3, 95%CI: 2.1-18.7. Reasons mentioned by respondents for missed doses included forgetting, alcohol use, poor access to care, work commitments, lack of food, stress and travelling. Of the 112 respondents in the study, 86.6% had unsuppressed viral loads.

**Conclusion:** Recommendations includes reinforcing use of reminders, automated SMS, establishing treatment supporters and collaborative efforts in reducing active substance use to improve patient adherence.

**Keywords:** Adherence; Adult; Antiretroviral (ARV) treatment; Health Belief Model; Human Immuno-deficiency Virus (HIV); Non-adherence.

#### Background

Namibia's Anti-Retroviral Therapy (ART) roll out has been very successful, achieving more than 84% coverage against a national target of 90% [1]. However, despite the successful role out of ART in Namibia, patients often fail to meet the level of adherence required for successful ART. The challenge of non-adherence to ART treatment has also been noted at the study site. In a study done in Nigeria, outcomes of non-adherence to ARV medication included viral resistance, treatment failure, toxicities and waste of financial resources [2]. Results from a study conducted in Botswana concluded that there is a strong correlation between adherence and clinical outcomes [3]. Given the large number of patients whose HIV infection will progress to AIDS if adherence is suboptimal, research is urgently needed to determine factors influencing adherence so that the most effective interventions to ensure adherence in African cohorts can be employed [4]. In a study in Ethiopia, 26.5% of the respondents had sub optimal adherence [5]. Locally, a recent study from Northern Namibia reported that 22% of respondents were non-adherent measured by pill count assessment. Henceforth this study envisaged to identify factors associated with non-adherence to ARV treatment in adult patients at Keetmanshoop hospital in Namibia.

Focusing on the study site, in January 2015, the Keetmanshoop Hospital had 1289 adults receiving ART. On average, the clinic serves about 250 to 400 patients in a month. In the period October to December 2014 a total of 933 patients received follow-up care at the study site. Of these patients 53% (n=530) achieved required adherence scores of more than or equal to 95% while approximately 47% (n=403) achieved less than the required 95%. The suboptimal adherence rate in 47% of the patients 36.7% found in January 2015 at the Keetmanshoop Hospital was significantly higher than that reported in the Northern parts of the country. The Health Belief Model (HBM) was the theory of choice in this study due to its applicability to patient adherence and preventative health practices. The HBM has four major components [6,7].

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## Perceived susceptibility

The assumption of the study was that patients whose perception of their susceptibility of HIV disease progression to AIDS is suboptimal may lead to non-adherence to ARV treatment.

## Perceived severity

These personal views of the HIV positive adult that the HIV diagnosis is a serious diagnosis with severe implications to health therefore requires appropriate action like adhering to medicines. This concept has a direct bearing on how the patient will adhere to the ARV treatment. The assumption of the study was that the lower the perception of severity the higher the non-adherence to ARV treatment.

## Perceived benefits and Costs

The individual patient needs to believe that adherence to ART would reduce susceptibility to HIV disease progression and severity. This was the basis for questions in the questionnaire which inquired about how essential ART was to the individual patient. The perceived cost shows that the individual's belief that the materials, physical and psychological costs of adhering to ART are all outweighed the benefits.

## Motivation

This component stresses that even if all components are in place, an individual motivation to take ARVs as prescribed is necessary, absence of which may result in non-adherence. This is influenced by availability of social support (e.g., family members).

## Enabling or modifying factors

The modifying factors include patient satisfaction and socio-demographic factors

## Research Methods

A combination of quantitative and qualitative was used to identify the adherence of participants and investigate the factors associated with non-adherence respectively. Numerical data was used in calculating patient adherence using pill count method and regression analysis on identifying factors statistically significant to non-adherence. Qualitative approach was used in form of a questionnaire to solicit mainly patient reasons for missed doses and their individual comments about service and taking ART.

## Population

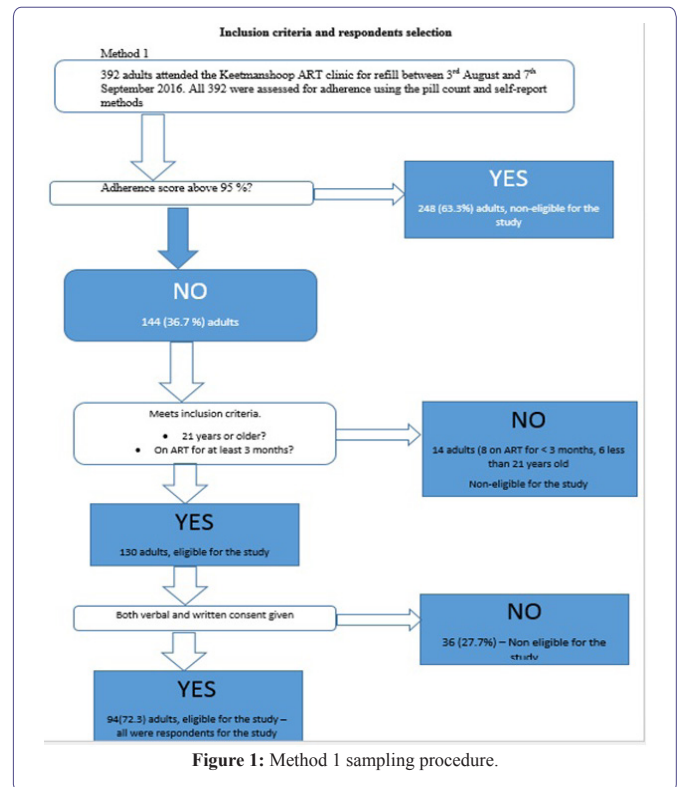
Population referred to HIV positive adults receiving HIV care at the Keetmanshoop Hospital. As of January 2015 the total number of adults in care at the hospital was 3568, which included 1289 ART people and 2279 pre-ART people.

## Target population

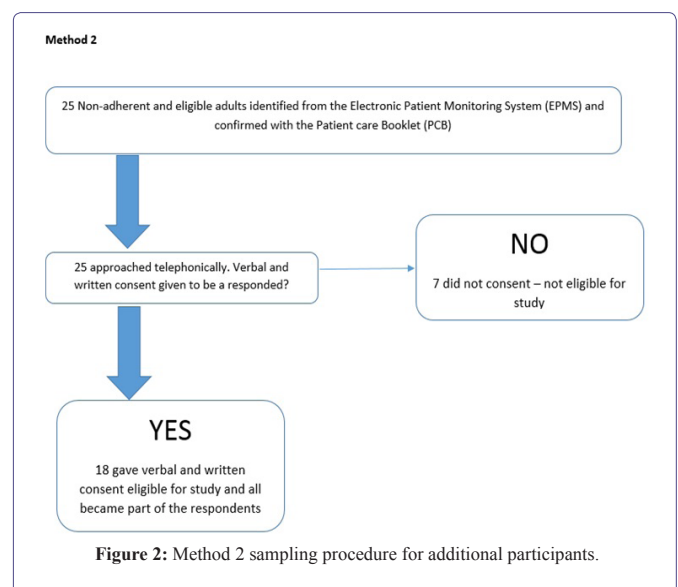
The target population were HIV positive adults 21 years and older on anti-retroviral treatment at Keetmanshoop Hospital. According to the EPMS of the 1289 patients who were on ART at the site. In the period October to December 2014, 439 patients (47%) out of 933 who attended the hospital were assessed as being non-adherent to their ARVs as their adherence score according to pill count was less than 95%.

## Sampling methods

Census and multi-stage sampling method was used as shown in the flow chart below in figure 1: Method1sampling procedure. Data collection period was from 3rd August to 7th September 2011.



The sample was insufficient and needed additional participants. These were added using review of records method as indicated in the flow chart below figure 2.



Altogether 112 adults gave a written consent to participate in the study and these made up the study sample.

## Data collection approach and method

Data collection was through structured interviews supplemented with review of patient records. Information reviewed in the records included the attendances of appointments dates, adherence assessment scores where non-adherence would be identified, ARV regimen of the particular patient, duration on ARV treatment and blood results of patients including the patient viral load.

## Development and testing of the data collection instrument

Whilst the patient adherence was calculated using the pill count and self-report methods, the questionnaire was compiled to investigate factors associated with non-adherence. Data from literature on factors related to non-adherence together with the Namibia HIV Patient Care Booklet (PCB) which is the manual tool in which patient information is entered on every follow-up visit to the clinic were used in questionnaire compilation. The tool was valid in Namibian context and reliable with a Cronbach alpha of 0.80. The questionnaire had 5 subsections; Socio-demographic and socio-economic, Patient-related factors, System-related factors, Disease- and treatment-related factors and Cultural and religious factors.

## Data collection process

Participants were allowed to choose their preferred venue for data collection, 73.3% (n=82) of respondents were interviewed at the clinic while 26.7% were interviewed at places of their choice other than the clinic. Each interview took approximately 10-15 minutes.

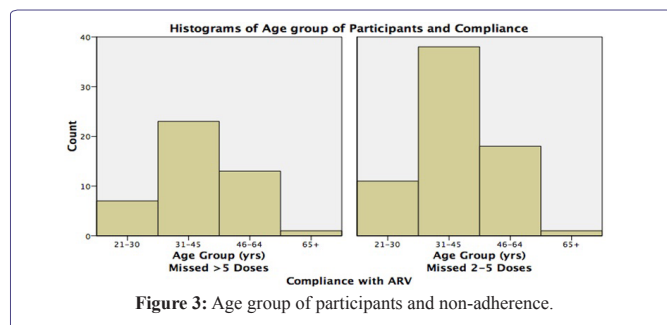
## Data analysis

Data was entered in SPSS version 21 and the data was checked for errors and prepared for analysis. Frequencies and graphs were generated. Correlation of variables was done to remove multicollinearity. Stata version 12 was used for logistic regression to do univariate and multivariate analysis to determine factors associated with missing five or more doses.

## Results

The table 1 below summarizes the socio-demographics of the sample.

The analysis of age group and non-adherence of respondents is shown below in figure 3.



**Figure 3:** Age group of participants and non-adherence.

The outcomes of correlation of variables using logistic regression to do univariate and multivariate analysis are shown below in table 2.

Patient Characteristics	Coding Categories	Number (%)
Sex	Male	49 (43.8%)
	Female	63 (56.3%)
Age in completed years	21-30	18 (16.1%)
	31-45	61 (54.5%)
	46-64	31 (27.7%)
	65 and above	2 (1.8%)
Ethnicity	Nama-Damara	53 (47.3%)
	Oshiwambo	53 (47.3%)
	Caprivan/Okavango	5 (4.5%)
	Otjiherero	1 (0.9%)
Religion	Catholic	30 (26.7%)
	Orthodox	63 (56.3%)
	Protestant	19 (17%)
	Islam	0
	Hindu	0
	African tradition	0
Marital status	Unmarried	80 (71.4%)
	Married	25 (22.3%)
	Separated/Divorced	5 (4.5%)
	Widow or widower	2 (1.8%)
Level of education	Illiterate	6 (5.4%)
	Basic education	73 (65.2%)
	Elementary education	5 (4.5%)
	College diploma and above	2 (1.8%)
Living condition	Living alone	16 (14.3%)
	Living with family	79 (70.5%)
	Living with friend	6 (5.4%)
	Living with others	11 (9.8%)
Source of support	Self-support	65 (58%)
	Families	36 (32.1%)
	NGOS	7 (6.3%)
	No support	4 (3.6%)
Current employment status	Employed	55 (49.1%)
	Unemployed	57 (50.9%)

**Table 1:** Socio-demographic and economic characteristics of patients (n=112).

In the univariate analysis the following factors were associated with missing five or more doses: Viral load OR: 5.6, 95%CI: 2.0-16.0, Employment OR: 3.3, 95%CI: 1.5-7.2, Taking active substance OR: 2.5, 95%CI: 1.3-4.7, and Emotional status OR: 1.6, 95%CI: 1.02-2.6. The ART stocks seemed adequate as the OR for out of stock did not lead to missed doses (OR: 0.6, 95%CI: 0.4-0.9). Factors such as gender, education, religion and marital status were not associated with missing five or more doses. After multivariate logistic regression the following factors were associated with missing five or more doses; Taking substance adjusted OR: 3.3, 95%CI: 1.4-7.6; WHO clinical stages 2 and 3 adjusted ORs and 95%CI: 4.1 (1.2-13.8) and 6.1 (1.5-25.1) respectively and viral load adjusted OR: 6.3, 95%CI: 2.1-18.7. Participants taking of active substances and the non-adherence were analyzed and the findings are demonstrated in figure 4 below.

Figure 3 (above) compares missed doses by taking active substances and gender. More women took active substances compared to men and thus were more likely to miss more than five doses compared

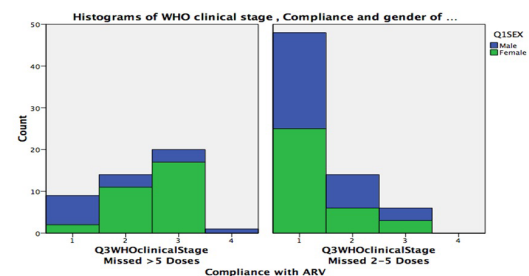
to men. Those patients who were self and family supported were more likely to miss their doses compared to those supported by NGOs and those who had no support. The participants' viral load and non-adherence correlation analysis results are illustrated in figure 5 below.

Variable	OR	95% CI
Gender	2.142	0.970-4.734
Age group*	1.104	0.642- 1.909
Ethnicity	0.673	0.361-1.252
Religion	0.601	0.330-1.095
Marital status	1.281	0.723-2.271
Education	0.989	0.574-1.075
Living conditions	0.835	0.500-1.394
Viral load	5.656	2.00-15.96*
Employment	3.254	1.463-7.235*
ART essential	0.931	0.482-1.797
Comfortable taking ART	0.842	0.563-1.260
Active substance	2.484	1.142-5.405*
Takes substance	2.516	1.338-4.730*
Disclose to community	1.855	0.783-4.395
Emotional status	1.623	1.022-2.578*
ARV remind HIV	1.356	0.788-2.334
Convenient clinic	1.576	0.964-2.576
Satisfied with care	1.08	0.608-1.920
Out of stock	0.588	0.3980-0.869*
Experience side effects	0.714	0.391-1.805
ARV and food	1.473	0.687-3.157

**Table 2:** Correlation of variables and non-adherence.

**Key:** \*Statistically significant

More females had viral load of more than 1000 copies per ml and missed more females missed more than 5 doses compared to males. Figure 6 below shows the outcomes of correlational analysis of participants WHO clinical stage and non-adherence.



**Figure 6:** Histograms of WHO clinical stage, compliance and gender.

### Respondents' responses on the reason/s for ARV missed dose (N=142, Total=100%)

All the 112 respondents in the study were asked for the reasons for missed dose/s. Some respondents mentioned more than one reason. The total number of reasons mentioned by the 112 respondents was 142. However, some reasons were recurring among respondents and were summarized together. Reasons mentioned in the order of most common to least commonly were; forgetfulness (21.8%), alcohol use (15.5%), challenges in accessing treatment (9.9%), work commitments (9.9%), transport problems to health facility (7%), lack of food (4.9%), stress (4.2%) and travelling (4.2%). A notable 4.2% of the respondents mentioned discomfort due to ARVs including the side effects as reason for missed dose.

## Discussion

### The level of non-adherence at the study site

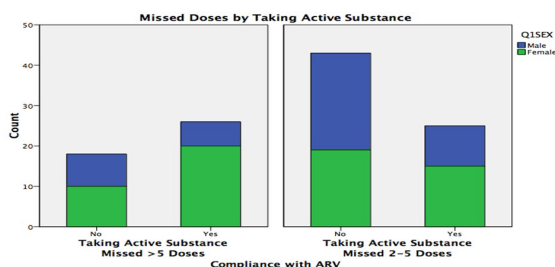
This study concluded that there was an abnormally high non-adherence at the study site warranting corrective interventions. The non-adherence was correlating with a very high number of patients with unsuppressed viral loads. The study results concur with those in another study that showed viral loads are strongly dependent on patient adherence [3].

### Factors associated with non-adherence

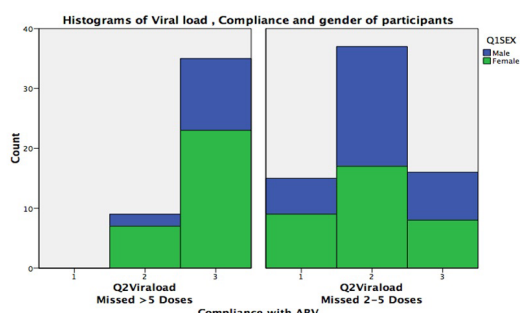
The study findings are discussed below under the five subsections of the data collection tool.

### Socio-demographic and economic characteristics

Firstly, the age group of 31-45 was found to be statistically associated with non-adherence. Possibly because most adults in this age group are actively employed which could have increased their chances of missing doses through work commitments, being away from home or being unable to collect their medicines from the clinic. On the contrary, a study in Ethiopia found no association between age and non-adherence [8]. Secondly, employment status also yielded statistical significance with regard to non-adherence. Although an Ethiopian study reported that the likelihood of ART non-adherence in employed patients was 0.41 times that of the unemployed patients, 5 in Northern Namibia employed respondents had a higher rate of adherence



**Figure 4:** Use of active substances and non-adherence.



**Figure 5:** Viral load and non-adherence.



compared to pensioners and unemployed respondents combined [9]. Gender and educational status did not yield any significance whilst men were more non-adherent than women in the North of the country [9]. Other factors which were statistically insignificant in this study were; ethnicity, religion, marital status, level of education, living condition and source of support. Although in some studies association between marital status and adherence levels were also not established, in Zambia unmarried people were found more non-adherent [8,10,11]. Similarly, some studies did not yield any association between education level and adherence levels, but others have documented significance [8,10,12,13]. Pertaining to source of support, it is highlighted that there was a significant association between availability of social support in taking ARV treatment and adherence to treatment in Ethiopia [8].

### Patient-related factors

Firstly, active substance abuse came out as a statistically significant factor associated with non-adherence. A little under half of the respondents were into active substances use and 43% of substance users highlighted they had missed an ARV dose at least once due to the substance use. This is almost double that reported in Northern Namibia where only 22.5% in the study respondents used active substances [9]. Secondly, patient emotional status also had statistical significance in relation to non-adherence. This study results show that 22% of the non-adherent adults were depressed while 6 (4.2%) respondents singled stress as a reason for missing ARV doses. Other studies have also concurred that depression is associated with non-adherence [8]. The association between emotional status particularly depression and non-adherence is further re-iterated elsewhere [10].

Factors which were insignificant to non-adherence included; Patient belief that ART is essential for their HIV care, comfortable to take ART in the presence of others, community disclosure and whether the patient feel taking your ARVs reminds them of the HIV infection. On the other hand, it has been recorded that lack of community disclosure is associated with non-adherence [14]. In addition, Gari et al., have also reported that poor beliefs in the benefits of ARVs were associated with non-adherence [15].

### System related factors

The ART stocks were statistically adequate. This meets the Namibia National Drug Policy mandate for the constant availability of safe and efficacious medicines across the Namibian population [16]. Similar findings are reported by Abdissa where the majority (94.2%) of the respondents had not encountered problems in refilling their ARV drugs at the pharmacy [8]. On the contrary, in Nigeria Uzochukwu et al., reported that non-availability of drugs at treatment sites was one of the factors related to non-adherence [2]. Although, in this study only 50% of the respondents found the clinic convenient for their care, this did not yield any statistical significance. The remaining two system related factors (convenience of clinic and satisfaction with health care workers) investigated in this study were also statistically insignificant. However, other studies have revealed a significant relationship between perception of study respondents on patient-healthcare worker relationship and their adherence to antiretroviral treatment [8,17].

Additionally, respondents cited some system related barriers as reason for missed doses. These barriers included transportation costs to the clinic, unfriendly service hours and high healthcare worker-patient

ratios and the resultant long waiting hours. Similar findings have been reported in Uganda, where transportation costs accounted for 7.8% of reasons why patients were non-adherent [18]. The association between transport challenges and non-adherence were also reported from other studies [19,20]. As reported in the Presidential Commission of Enquiry, the public health sector suffers from inadequate numbers of staff and availability of equipment and resources [21].

### Disease and treatment related factors

Under this subsection all the four aspects investigated (duration on ARV treatment, experience of any side effects to ARVs, special instruction regarding ARVs and food and general health status in past 1 month) they were statistically insignificant. Whilst other reports note that taking ART for more than two years was positively associated with adherence in high income countries, others have not yielded any statistically significant association between duration on treatment and adherence to ART [8,15,22]. Regarding ARV side effects, they range from minor ones such as nausea, vomiting, skin rash to severe liver damage or life threatening Steven Johnson Syndrome [1]. In this research it was worrisome to note that 57.1% of respondents were having or had previously suffered from ARV side effects. Uzochukwu et al., also documents physical side effects of ARVs among reasons for non-adherence [2]. Although only 53.6% of respondents had special instructions regarding ARVs and food, the majority of respondents believed that they could not take their ARVs when they have not eaten. This belief is contrary to Namibian ART guideline which discourages patients from missing their ARVs because they do not have any food [1]. 4.9% of this study respondents commented that not having food to take with the ARVs as the reason for missed dose. These findings are similar to those reported by Nghoshi where 7% of study respondents complained of dietary requirements [9].

While non-adherent respondents would be expected to be sick and with higher WHO clinical stage, only 20% of the respondents reported being sick in the past one month. The results of non-adherence are cumulative over time, thus respondents who were non-adherent during the data collection period might still have been healthy, but with continual non-adherence, a depleted immune system and rising risk of opportunistic infections would eventually become sick patients with time.

### Cultural and religious beliefs related factors

Similarly all four factors investigated in this section were found to be insignificant statistically. These were; whether taking ARV medicines offend one's cultural beliefs, if HIV infection can be treated by cultural methods without ARV medicines, whether taking ARV medicines offend any of your religious beliefs and if HIV infection can be treated by religious methods without ARV medicines. In a study in Northern Namibian, 11% of respondents reported taking other traditional treatments together with ART [9]. It is noteworthy that a South African study reports that there is often tension between ARV treatment and alternative therapies [23]. Such beliefs present healthcare workers with a challenge to promote and foster adherence to ART among patients. In an era where some churches offer miracles of different forms, it appears HIV patients have not been spared and are sometimes advised to stop taking the ARVs as a sign of their healing resulting in non-adherence of the part of the patients.

## Respondent's last viral load

After multivariate logistic regression viral load was statistically related to non-adherence. It is noteworthy that 86.6% respondents in this study had unsuppressed viral load while only 13.4% had non-detectable viral loads, also known as maximum viral suppression. On the contrary, in a study in Myanmar, 94% of patients had an undetectable viral load and only 4% had ART failure. It is further reported that 84.7% of respondents on first and second line treatment in the Myanmar study had viral load below 1000 copies/ml [24]. The viral loads in this study support the pill count adherence method as the viral loads depict the adherence level in patients. This is a public threat as resistant HIV strains could result due to the non-adherence [3].

## Respondents WHO clinical stage

Multivariate logistic regression yielded statistical significance in both WHO clinical stages 2 and 3. As noted above, respondents' non-adherence may not have had immediate negative outcomes but the consequences of continual non-adherence are likely to be more prominent over time. In this study, the respondents viral load were already showing non-adherence although the majority of the patients were still in stage 2.

## Respondents' responses on the reason/s for ARV missed dose (N=142, Total=100%)

The majority of the reasons were patient related reasons, for example alcohol use, followed by socio-demographic, system of care, disease and treatment and lastly religious reasons. Some of the reasons mentioned in this study, for example forgetfulness, transport problems and work commitments are similar to those reported in other studies [5,8,9]. However, while lack of food was mentioned in this study as reason for missed dose, in Northern Namibia, lack of food was not revealed as a reason for missed doses [9]. Due to the Savanna Climate in Northern Namibia, crop farming of staple foods like sorghum is possible which helps to enhance food security in most households. In contrast southern Namibia is predominantly semi-arid to desert which does not support crop farming but mainly small livestock husbandry [25]. Of note also is that, more than half of the respondents in this study were unemployed, which may have impacted on food insecurity among some respondents. As the study site serves a very vast geographic area, is also not surprising that some patients found it very difficult accessing the health facility. A Nigerian study reports that HIV infection related sicknesses constituted 3.5% of reasons for missed dose, while 2.1% simply got tired of taking ARVs and missed their doses [2].

## Respondents' comments

The majority of the respondents' comments were directed towards the service delivery system. Main themes derived from the comments included; patients' satisfaction with the services they received from the nurses; although acknowledging inadequacy of nursing staff. Some respondents expressed the need for more nurses to be allocated to the division. Another commonly mentioned suggestion was need for decentralization of ART services so that service can be nearer to the people. Other respondents suggested that the duration of ARV prescription should be increased for up to 6 months to eliminate frequent visits the clinic for refill. Another noteworthy suggestion was towards the integration of ART with other services, including the dispensing of medicines and reviews by a medical officer.

## Study recommendations

The interpretation of the research findings were presented in relation to the Health Belief Model theory and the conceptual framework of the study on factors associated with non-adherence. The HBM theory used in this study assumed that the patient's ability to adhere to ART was strongly dependent on a number of concepts.

**Perceived severity:** Healthcare workers should use results of viral loads to explain resistance at appropriate levels of understanding of patients as a way to promote understanding of the consequences of non-adherence.

### Perceived benefits:

- Healthcare workers are advised to plot patients' biological markers including CD4 and viral loads on a graph to explain benefits of adherence to patients
- Recruitment and the active involvement of "expert patients" these are HIV positive patients who have disclosed their HIV status and trained to motivate other patients and explaining benefits of adhering to treatment and practicing healthy lifestyle while on ART treatment

### Perceived barriers:

- A multidisciplinary approach is required to help patients to reduce or stop active substance use
- Healthcare workers need to strengthen patients' referral system across the different levels of HIV care so as to make it easier for patients to access continuing care when travelling
- To improve access to ART services for patients in remote areas, ART can be integrated into existing outreach programmes
- Initiate and strengthen effective HIV workplace programs so as to increase awareness among employers on the need for treatment support of employees as well as to reduce stigma which is often experienced at workplaces
- The MoHSS needs to strengthen collaborative efforts with other non-governmental organizations on supportive care for ARV patients such as by providing food support
- Lay counsellors need to be trained in basic communication skills and counselling techniques

### Cues to action:

- Patients should be encouraged to identify treatment supporters of their choice
- The MoHSS is recommended to adopt the use of Automated Short Message Service (SMS) as a way of reminding patients their ARVs Self-efficacy
- Healthcare workers should strengthen health education how to manage in case of missed doses
- There is need to anticipate and discuss potential side effects with patients, the time they maybe experienced and importantly how to handle them

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## Author Contributions

BLD contributed to the study design particularly the sampling method and data collection method. Both authors revised the manuscript, gave final approval to the version published and they agree to be accountable for all aspects of the article.

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