Validity of World Health Organisation prescribing indicators in Namibia's primary health care: findings and implications

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Abstract

Objective: World Health Organization/ International Network of Rational use of Drugs (WHO/INRUD) indicators are widely used to assess medicine use. However, there is limited evidence on their validity in Namibia's primary health care (PHC) to assess the quality of prescribing. Consequently, our aim was to address this. Design, setting, participants and interventions: An analytical cross-sectional survey design was used to examine and validate WHO/INRUD indicators in outpatient units of two PHC facilities and one hospital from 1st February 2015 to 31st July 2015. The validity of the indicators was determined using two-by-two tables against compliance to the Namibian standard treatment guidelines (NSTG). The receiver operator characteristics for the WHO/INRUD indicators were plotted to determine their accuracy as predictors of compliance to agreed standards. A multivariate logistic model was constructed to independently determine the prediction of each indicator. Main outcomes and results: Out of 1243 prescriptions; compliance to NSTG prescribing in PHCs was sub-optimal (target was >80%). Three of the four WHO/INRUD indicators did not meet Namibian or WHO targets: antibiotic prescribing, average number of medicines per prescription and generic prescribing. The majority of the indicators had low sensitivity and/or specificity. All WHO/INRUD indicators had poor accuracy in predicting rational prescribing. The antibiotic prescribing indicator was the only covariate that was a significant independent risk factor for compliance to NSTGs. Conclusion: WHO/INRUD indicators showed poor accuracy in assessing prescribing practices in PHCs in Namibia. There is need for appropriate models and/or criteria to optimize medicine use in PHCs in the future.

Introduction

The World Health Organisation (WHO) estimates the global burden of inappropriate prescribing of medicines to be over 50% [1,2]. This is important especially in low and middle income countries (LMICs) which have the highest burden of non-communicable and infectious diseases. Moreover, the cost of medicines can account for up to 70% of total healthcare costs in LMICs, with a significant proportion out-

of-pocket [2-4]. Consequently, it is very important to optimize medicine use in these settings. In Namibia, over 45% of the adult population currently live with hypertension [5], with cardiovascular diseases (CVD) currently accounting for 21% of annual deaths [6]. There are also high rates of infectious disease in Namibia leading at times to excessive and over use of antibiotics [7,8]. As a result, there is a need for robust systems and indicators to monitor and guide medicine use to enhance appropriate prescribing.

The WHO/ International Network for Rational Use of Drugs (WHO/INRUD) prescribing indicator framework was developed as an objective measure for evaluating the appropriateness of medicine use [2,9,10]. These indicators are currently seen a proxies for assessing the quality of prescribing as currently no international standards have been empirically developed [2]. This is a concern as these indicators are widely integrated into health management information systems (HMIS) across countries [2,11,12], and no indicator directly measures rational prescribing according to the WHO definition [13].

Namibia's public health care is based on a Primary Health Care (PHC) system, funded mainly by the government. Currently, the WHO/INRUD indicators are integrated into the Namibian health system at all levels of healthcare [13]. The current prescribing targets and rates for the five key indicators are [2,9,14-17]:

- Average number of medicines per encounter WHO Target; Namibia target <2, acceptable 2.5; currently in Namibia: 1.6 3.1. However, a high number of medicines per prescription may not necessarily mean irrational prescribing as seen in patients with chronic diseases or multimorbidities [9,18]
- % of encounters with an antibiotic prescribed WHO Target <30%, Namibia target <25%, acceptable 35%; currently in Namibia: 56 80%. It is difficult to assess whether these targets represent quality prescribing without knowing the presenting infections, which could be viral in origin as typically seen with upper respiratory tract infections (URTIs) [19] or the extent of underlying infectious diseases such as HIV and TB
- % of encounters with an injection: WHO Target <20%, Namibia target <10%, acceptable 15%; safe medication use in Namibia 44% 50%
- % of medicines prescribed by International non-proprietary name (INN) WHO Target 100%, Namibia target - 100%, acceptable – 80%; currently in Namibia: 74 - 80%. However, 100% may be difficult to achieve with substitution discouraged in a small group of medicines including those for epilepsy [20,21], and where there are concerns with the quality of generics [22,23]
- Compliance to Essential Medicine Lists WHO Target 100%. In Namibia compliance to Standard Treatment Guidelines currently at 15.4 – 44.6 % depending on the region; acceptable > 80%

The task to reliably measure the quality of prescribing in PHCs should compare with robust methodologies that have been used in other circumstances to assess the quality of care including the Integrated Management of Childhood Illness (IMCI) project, the Adult Primary Care Assessment Tool (PACT) and the quality indicators that have been developed for antibiotic prescribing [24-29].

Consequently the aim of this study was to appraise the applicability of current WHO/INRUD prescribing indicators in assessing the quality of prescribing in PHC settings in Namibia. Subsequently, use the findings if pertinent to suggest the development of different indicators to guide future prescribing. This includes compliance to Namibian Standard Treatment Guidelines (NSTG) [30]. The NSTG, which include treatment guidance for patients in both hospital and ambulatory care including community infectious diseases, are seen as appropriate for assessing the quality of prescribing based on the WHO definition of rational medicine use [12]. This is because STGs in LMICs are typically based on the principles of rational use of medicines and adapted from WHO recommendations. We are aware that different treatment guidance documents exist in Nambia, which can give conflicting advise [31]. Adopting the NSTG as the gold standard helps to address such concerns.

Methods

Study design and setting

A cross-sectional design was undertaken to determine the quality of medicine prescribing among public health care facilities in Namibia [9]. Prescribing habits of physicians in out-patient departments (OPD),

stratified by WHO/ INRUD prescribing indicators , were assessed against compliance with current NSTG [30].

The study was conducted in three OPD settings among public health facilities in Khomas region, one of the fourteen regions of Namibia that houses Windhoek, the capital city, with a cosmopolitan population [32]. The health facilities were selected based on the level of heath care provided, which included a clinic, health center and hospital settings, in line with WHO recommendations [9]. These facilities were purposively selected because of their large and diverse patient populations and, close proximity to each other. In addition, reflective of the situation across Namibia.

Study subjects and procedure

Patients at the OPD units were stratified by the level of care facility to include two PHC facilities – Khomasdal Clinic (KMDC) and Katutura Health Centre (KHC), and one hospital – Intermediate Hospital Katutura (IHK). The Khomasdal clinic was purposely selected out of the ten clinics in the Khomas region based on its proximity and demographic and service similarity to KHC and IHK. All patients recieving OPD care at the selected health facilities were included in the study. The sample size of 1243 for patient prescriptions was determined using the Kish Leslie method [33]. This estimation was based on an earlier study where compliance to STGs was 26%, giving a p = 0.26 [17]; with β power set at 80%, $\alpha = 0.05$ with a critical value of 1.96 for a two-tailed test.

To evaluate current prescribing practices, we evaluated the most recent prescription in the patients' health passport – that is a prescription written on the day of the study visit that may include more than one medicine. The data collection tool (Annex 1) was pre-tested among 10 patients at the OPD at IHK, and subsequently refined to improve its robustness. Prescribing data was collected from the patient health passports by the research team led by QN and three trained and experienced data collectors using the standardized data abstraction tool.

Inclusion and exclusion criteria

We included all out-patient prescriptions generated from 1st February 2015 to 31st July 2015. We excluded prescriptions from inpatient settings and specialist OPDs where the medicine use patterns are more complex. We also excluded patients where the diagnosis was not supported by the signs and symproms recorded in the patient's health passport and those on a follow up visit or reattendance.

Measurement, analysis, validation and criterion development

The WHO/INRUD prescribing indicators were used as predictors for compliance to the NSTG as the principal outcome measure. We determined the level of compliance and prescribing indicators in the OPD setting. The sensitivity and specificity of each WHO/INRUD prescribing indicator, e.g. the average number of medicines, % precriptions with antibiotics or injections and % of medicines prescribed by INN, was assessed as predictors for compliance to the NSTG to determine their validity. The association of the other covariates such as the level of health facility, patient demographics, medication prescribed, primary diagnosis and/or comorbidities, were also assessed against the WHO/INRUD prescribing indicators.

Quantitative data analysis was undertaken using SPSS Version 23. The WHO/INRUD prescribing indicators were determined using descriptive statistics of frequencies. The specificity and sensitivity of the prescribing indicators were determined by constructing two-by-two tables for each indicator. A plot of the sensitivity against the (1- specificity) was constructed to generate receiver operator characteristic (ROC) for each prescribing indicator. The area under the curve (AUC) for the ROC was subsequently determined to derive the optimal specificity and sensitivity of the prescribing indicators in determing compliance to the NSTG.

The accuracy of the indicator in predicting compliance to the NSTG was classified according to the AUC point system: 0.90-1 = excellent, 0.80-.90 = good, 0.70-0.80 = fair; 0.60-0.70 = poor and <math>0.50-0.59 = fail. An indicator with an AUC value greater than 0.5 does not rely on chance, and can discriminate between prescriptions that are compliant or non-compliant to the NSTG [34]. Consequently, the AUC cut-off was set at 0.6 for a valid prescribing indicator. A level of AUC ≥ 0.6 is considered to be of satisfactory accuracy in evaluating the sensitivity and specificity of the indicator [34].

Compliance to the NSTG was determined using a correctness score sheet derived from a panel incorporating three experts – one clinical pharmacist, one consultant physician and QN from the research team. The association between the prescribing indicators and the compliance level was determined using bivariate analysis with the Chi-squared test. A multivariate logistic regression analysis was used to determine the odds ratios for each prescribing indicator in predicting compliance to NSTG prescribing. A student's T-test and/or ANOVA were used to determine associations between continuous variables. In this study, the level of significance (α) was set at *p* = 0.05 at a 95% confidence interval.

Ethics

Permission to conduct the research was granted by the University of Namibia (UNAM) and the Ministry of Health and Social Services (MoHSS). Specific patient and prescriber identifiers such as the names and ID numbers were not collected but rather a specific numbering was assigned to each study participant for purposes of identification.

Results

We will first discuss compliance to the NSTG and factors involved before discussing the characteristics of the prescriptions, in line with the principal objectives.

Level of compliance to the NSTG

Overall, out of the 1,243 prescriptions (one each from 1243 patients), the majority (73%) complied with the NSTG through the appropriate choice of medicine(s) or treatment for the diagnosis indicated. Compliance to treatment suggestions in the NSTG was significantly higher at PHC facilities (76.1%) than at the hospital (70.5%, p=0.03) (Table 1).

Characteristics		Compliand	ce to NSTG		
		Yes (%)	No (%)	P -value	Cramer-V
Facility level					
	Hospital	491(70.5)	205(29.5)		
	PHC	416(76.1)	131(23.9)	0.030*	0.062
Prescriber type	Madical officer		200(27.6)		
		765(72.4) 112(77.2)	36(22.8)	0 198	0.036
Patient gender	Nuise	112(77.2)	50(22.0)	0.150	0.050
Server	Female	505(73.1)	186(26.9)		
	Male	402(72.8)	150(27.2)	0.919	0.003
Patient age					
Ac	dult (≥ 16 years)	701(74.6)	239(25.4)		
Cl	hild (< 16 years)	206(68)	97(32)	0.025*	0.064
Antimicrobial used	Ň		200(24.5)		
	Yes	565(65.5)	298(34.5)	0.000*	0.254
Analgosic used	NO	342(90)	38(10)	0.000	0.254
Analgesic useu	Yes	512(72)	199(28)		
	No	395(74.2)	137(25.8)	0.380	0.025
Trained on STG		, , , , , , , , , , , , , , , , , , ,	,		
	Yes	291(72.2)	112(27.8)		
	No	616(73.3)	224(26.7)	0.676	0.012
Resp diagnosis					
	Yes	226(70.4)	95(29.6)	0.000	0.000
CV/E diagnosis	NO	676(73.8)	240(26.2)	0.239	0.033
CV3 ulagnosis	Vas	17(81)	<i>A</i> (19)		
	No	890(72.8)	332(27.2)	0 406	0 024
GIT diagnosis		030(7210)	552(2712)	01100	0102 1
U U	Yes	25(75.8)	8(24.2)		
	No	882(72.9)	328(27.1)	0.712	0.010
Comorbidity					
	Yes	106(75.2)	35(24.8)		
	No	801(72.7)	301(27.3)	0.531	0.018
Antibiotic prescribe	ea Voc	102(62 6)	276(26 1)		
	No	485(05.0)	60(12.4)	0 000*	0 263
Generic prescribing	1	12 ((0) 10)	00(12.1)	0.000	0.200
	, >80%	346(76)	109(24)		
	< 80%	559(71.1)	227(28.9)	0.062	0.053
Injection prescribin	g				
	Yes	105(78.4)	29(21.6)		
	No	802(72.3)	307(27.7)	0.137	0.042
# medicine prescrib	Ded		220/25 1)		
	S medicines	000(74.9) 251(69 1)	220(25.1) 116(21.6)	0.010*	0.067
		231(00.4)	110(31.0)	0.013	0.007

Table 1 - Bivariate analysis of factors associated with the compliance to NSTG prescribing

Resp = respiratory; CVS = cardiovascular, GIT= gastrointestinal tract; * = (p < 0.05)-Statistically significant- Pearson Chi-square Test

Factors associated with compliance to the NSTG by treatment indication

Compliance with the NSTG was significantly associated (p < 0.05) with the health facility level and type (p=0.03), the patients' age (p=0.025), the prescribing of antimicrobial/antibiotics (p<0.001) and the average number of medicines per prescription (p=0.019) (Table 2). There was no significant association (p > 0.05) between compliance to the NSTG and prescriber cadre, patient gender, prescribing by INN and the presence of analgesics or antihistamines on the prescription (Table 1). There was also no association between the level of INN prescribing, the use of injections and prescriber type with compliance to the NSTG (Table 1).

Validity of the prescribing indicators against compliance to the NSTG

The sensitivity rates of the WHO/INRUD indicators ranged from 11.6% to 72.3% (Table 2). The average number of medicines per prescription had an acceptable sensitivity (72.3%, p =0.019). The specificity of the WHO/INRUD indicators ranged from 17.9% to 91.4% (Table 2). The injection prescribing indicator had an acceptable specificity of 91.43% (p =0.137) (Table 2).

	NSTG Compl	iant	Totals (%)	p-value	Cramer V	
	Yes (%)	No (%)				
Antibiotic indicator						
Positive (759)	483(53.3) [‡]	276(82.1)	759(61.1)	0.000*	0.263	
Negative(484)	424(46.7)	60(17.9)^	484(38.9)			
Injection indicator						
Positive (759)	105 (11.6) [‡]	29(8.6)	134(10.8)	0.137	0.042	
Negative(484)	802(88.4.%)	307(91.4)^	1109(89.2)			
Generic indicator						
Positive (759)	346(38.2) [‡]	109(32.4)	455(36.7)	0.60	0.053	
Negative(484)	559(61.8%)	227(67.6)^	1109(63.3)			
Average medicines						
< 3 medicines (759)	656(72.3) [‡]	220(65.5)	876(70.5)	0.019*	0.067	
>3 medicines (484)	251(27.7)	116(34.5)^	367(29.5)			

Table 2. Bivariate analysis: validating prescribing indicator against compliance to NSTG

= sensitivity; ^ = Specificity; * = significant association - p < 0.005</pre>

ROC for the WHO/INRUD indicators against compliance to NSTG

The ROC test yield failed accuracy for three indicators: Injection prescribing (AUC=0.49; 95%CI: 0.45 - 0.52, p=0.421), average number of medicines per prescription (AUC=0.46; 95%CI: 0.43 - 0.5, p=0.045), generic prescribing (AUC=0.51; 95%CI: 0.48 - 0.55, p=0.46). The antibiotic prescribing indicator had a poor accuracy (AUC=0.49; 95%CI: 0.61 - 0.68, p=0.001) (Figure 1).



Figure 1 – ROC Curve for medicine use indicators

Multivariate logistic model for factors associated with compliance to the NSTG A logistic regression analysis was conducted to predict compliance to the NSTG for 1240 patients based on the health facility level, prescriber cadre category, patient demographics, diagnosis by body system and the category of medication prescribed as predictors (Table 3). Three prescription records were excluded from the logistic regression analysis due to one or more missing characteristics. A test of the full model against a constant only model was statistically significant, indicating that the predictors as a set reliably distinguished between compliance and non-compliance to the NSTG prescribing in the PHCs (χ^2 = 147.2, *p*<0.001 with *df* = 34). Nagelkerke's R² of 0.163 indicated a relationship between prediction and grouping by compliance to NSTGs. Prediction success overall was 73.4% (96.6% for compliance to the NSTG and 11.3% for non-compliance). The Wald criterion demonstrated that only prescribing of an antibiotic and/or antimicrobial (*p*<0.001) made a significant contribution to the prediction.

The WHO/INRUD indicators including the average number of medicines per prescription, INN prescribing rates and the extent of injection prescribing, were not significant predictors for compliance to the NSTG.

In addition, patient's demographics, prescriber cadre category, diagnosis and the health facility level were also not a significant predictor for compliance to the NSTG. One covariate - antibiotic/ antimicrobial prescribing (OR=0.2, 95%CI: 0.20 - 0.48) - was identified as an independent risk factor for compliance to the NSTG (Table 3). Prescribing of an antibiotic decreases compliance to the NSTG by 80%.

Table 3: Multivariate logistic model for compliance to NSTG prescribing

Covariates	OR(95% CI)	p -value
Facility level		
Hospital	0.7 (0.5, 1.1)	
PHC	1.0	0.114
Prescriber cadre		
Medical	0.7 (0.5, 1.1)	0.127
Nursing	1.0	
Completed STG training		0 50 4
Yes	1.1 (0.8, 1.6)	0.534
NO Detientie gender	1.0	
Fallent's gender	1 1(0 8 1 4)	0 677
Male	1.1(0.8, 1.4)	0.077
Patient's age (vrs)	1.0	
Adult (≥ 16 years)	0.9(0.6.1.2)	0.403
Child (< 16 years)	1.0	
Antibiotic prescribed		
Yes	0.2(0.2,0.48)	0.000*
No	1.0	
Injectable prescribed		
Yes	1.5 (0.9,2.4)	
No	1.0	0.122
Number medicines prescribed		
< 3	1.1 (0.8, 1.4)	0.714
> 3	1.0	
% generic medicines prescribed		
Yes	1.2 (0.9,1.6)	0.309
No	1.0	
Diagnosis by system		
Gastrointestinal	1.2 (0.7,2.0)	0.439
Musculoskeletal	1.5(0.9,2.5)	0.094
	0.4(0.1,1.2) 07(0223)	0.110
Obstetrics and gypaecology	0.7 (0.2, 2.3) 0 7 (0 2 2 1)	0.303
HEENT	15(0.2,2.1)	0.470
Lower Respiratory Tract	1.5(0.5,2.4) 1.5(0.54.8)	0.495
Genital Urinary Tract	1.1 (0.6.2.1)	0.721
Fever	0.9 (0.3,2.3)	0.821
Non specific diagnosis	1.0	
Comorbidity		
Yes	1.3 (0.8,2.0)	0.247
No	1.0	
Treatment prescribed		0.182

Antimicrobial	1.0	
GIT medication	0.6 (0.3, 1.0)	0.053
Decongestants	0.3 (0.1,1.1)	0.069
Antihistamines	0.6 (0.2,1.7)	0.350
Cardiovascular	3.5 (0.9,12.7)	0.063
Psychiatric	2.4 (0.6,9.9)	0.222
Emollients	1.2(0.2,6.0)	0.821
Non – pharmacotherapy	1.1(0.5,2.6)	0.845
Endocrine medicines	4.0(0.4,38.5)	0.226
Rehydration salts	0.6(0.3,1.2)	0.113
Anti-asthmatic agents	0.6(0.1,3.8)	0.618
Analgesics/antinflammatory	1.1(0.8,1.7)	0.501
Constant	8.1	0.000

Characteristics of patient prescriptions at OPD units

The majority of prescriptions were initiated at the OPD of the hospital (56%; p < 0.001), by physicians (87.3%; p < 0.001), for females (55.6%; p < 0.008) and adults (≥ 18 years - 73.4%). Table 4 shows the bivariate analysis of the WHO/INRUD indicators. The antibiotic prescribing indicator was significantly higher among nurse prescribers, children, prescriptions with an antimicrobial, a respiratory diagnosis and patients with comorbidities. Antibiotic prescribing was lower with a diagnosis of a cardiovascular or gastrointestinal condition. Other areas of interest included INN prescribing associated with the health facility (p<0.001), the type of prescriber (p=0.033), co-medication with analgesic (p<0.001) and prescriber training on STGs (p<0.001).

Characteristics	Prescription has an antibiotic prescribed		No. medicines per prescription			Prescription has an Injection			% INN prescribing/prescription			
	Yes (%)	No (%)	P - value	<3 (%)	> 3 (%)	P - value	Yes (%)	No (%)	P - value	≥80%	<80%	P - value
Facility level												
Hospital	409(58.8)	287(41.2)		456(65.5)	240(34.5)		89(12.8)	607(87.2)		145(20.9)	549(79.1)	
PHC	350(64)	197(36)	0.069	420(76.8)	127(23.2)	0.000	45(8.2)	502(91.8)	0.01	310(56.7)	237(43.3)	0.000
Prescriber type												
Medical officer	646(59.5)	439(40.5)		764(70.4)	321(29.6)		121(11.2)	964(88.8)		385(35.5)	698(64.5)	
Nurse	113(71.5)	45(28.5)	0.004	112(70.9)	46(29.1)	0.903	13(8.2)	145(91.8)	0.268	70(44.3)	88(55.7)	0.033
Patient gender	424(61.4)	2(7/20.0)		402(60.0)	200(20.1)		76(14)	(15(00)		257(27.2)	422/62.0	
Female	424(61.4)	267(38.6)	0.015	483(69.9)	208(30.1)	0 6 1 9	76(11) 58(10 5)	615(89) 404(80 F)	0 701	257(37.2)	433(62.8)	0.624
Detient and	555(00.7)	217(59.5)	0.813	393(71.2)	159(20.0)	0.018	56(10.5)	494(89.5)	0.781	198(55.9)	555(04.1)	0.034
Adult (> 16 years)	499(53.1)	441(46.9)		666(70.9)	274(29.1)		119(12 7)	821(87-3)		348(37.1)	591(62.9)	
Child (< 16 years)	260(85.8)	43(14.2)	0.000	210(69.3)	93(30.7)	0.609	15(5)	288(95)	0.000	107(35.4)	195(64.6)	0.609
Antimicrobial used		,		(,		()	()				
Yes	753(87.3)	110(12.7)		570(66)	293(34)		86(10)	777(90)		319(37)	544(63)	
No	6(1.6)	374(98.4)	0.000	306(80.5)	74(19.5)	0.000	48(12.6)	332(87.4)	0.163	136(36)	242(64)	0.740
Analgesic used												
Yes	433(60.9)	278(39.1)		540(75.9)	171(24.1)		83(11.7)	628(88.3)		225(31.6)	486(68.4)	
No	326(61.3)	206(38.7)	0.895	336(63.2)	196(36.8)	0.000	51(9.6)	481(90.4)	0.240	230(43.4)	300(56.6)	0.000
Trained on STG												
Yes	231(57.3)	172(42.7)		251(62.3)	152(37.7)		53(13.2)	350(86.8)		87(21.6)	316(78.4)	
No	528(62.9)	312(37.1)	0.061	625(74.4)	215(25.6)	0.000	81(9.6)	759(90.4)	0.062	368(43.9)	470(56.1)	0.000
Resp diagnosis	267(02.2)	54(46.0)		220(00.5)	101(21 5)		24(6.5)	200(02.5)		440(27.4)	202(62.0)	
Yes	267(83.2)	54(16.8)	0.000	220(68.5)	101(31.5)	0 202	21(6.5)	300(93.5)	0.004	119(37.1)	202(62.9)	0.966
CVC diagnosis	490(53.5)	420(40.5)	0.000	051(71.1)	205(28.9)	0.392	113(12.3)	803(87.7)	0.004	334(30.3)	580(03.5)	0.800
	2 (9 5)	19(90.5)		16(76.2)	5(23.8)		1(4.8)	20(95.2)		6(28.6)	15(71 /)	
No	2 (3.3) 757(61.9)	465(38.1)	0.000	876(70.5)	362(29.6)	0.563	133(10.9)	1089 (89.1)	0.370	449(36.8)	771(63.2)	0.438
GIT diagnosis	()	()			(3)		()	, (
Yes	14(42.4)	19(57.6)		25(75.8)	8(24.2)		1(3)	32(97)		13(40.6)	19(59.4)	
No	745(61.1)	465(38.4)	0.026	851(70.3)	359(29.5)	0.500	133(11)	1077(89)	0.146	442(36.6)	767(63.4)	0.638

Table 4 - Bivariate analysis of factors associated with the WHO/INRUD indicators

Comorbidity													
	Yes	99(70.2)	42(29.8)		70(49.6)	71(50.4)		10(7.1)	131(92.9)		48(34)	93(66)	
	No	660(59.9)	442(40.1)	0.018	806(73.1)	296(26.9)	0.000	124(11.3)	978(88.7)	0.134	407(37)	693(63)	0.493

r = (p < 0.05)-Statistically significant- Pearson Chi-squared Test; INN International non-proprietary name

WHO/INRUD prescribing indicators at the primary healthcare facilities

3,759 medicines were prescribed among the 1,243 prescriptions. The average number of medicines per prescription was 3.02 ± 1.14 (IQR = 2 - 4). This was highest at the hospital level (3.15 ± 1.197) compared to the clinic (2.93 ± 1.142) and health center (2.85 ± 1.017) (p=0.004). 63% of medicines were prescribed by INN and more than two thirds of the prescriptions included an antibiotic - 69% (; p = 0.059). The percentage of prescriptions with ≥ 80% of medicines prescribed by INN was 36.6% (n= 382/1241). Injection prescribing was 10.8% (n=134/1243), highest at the hospital 12.8%.

Discussion

The majority of the prescribing indicators according to the WHO/ INRUD criteria were sub-optimal among these facilities in Namibia (Table 4). However, 70% of prescriptions in this study had three or less medicines prescribed, with an average of 3.12 ±1.14 medicines per prescription. This is similar to other African countries [2]. 3759 medicines (63.9%) were prescribed by INN, which is also similar to other African countries [2] although lower than Ethiopia at 98.7% [35] and Botswana at 79% [36]. This needs to be addressed to avoid patient confusion if different branded generics with different names are dispensed on each occasion, with the drive towards lowering generic prices through increased competition [20], and patients are unsure whether they are taking the same medicine [37]. There are also concerns with the use of injections especially in hospital (Table 1). This will be investigated further to ensure injections are not given unnecessarily in the future.

Encouragingly, the majority (73%) of prescriptions were compliant to NSTG recommendations (Table 2). This rate is an improvement on a previous study conducted among public facilities in Namibia [17], and compares favourably with a recent study among PHCs in Botswana where there were concerns with antibiotic prescribing [36]. This high compliance rate in Namibia also compares favourably with developing countries where compliance rates have ranged from 30% to 50% [10,38]. One possible reason why our study found a higher compliance rate may be due to the fact that we excluded prescriptions where the diagnosis was not supported by the recorded signs and symptoms. Having said this, our compliance rate is lower than the compliance level set at 85% by MSH [39] and there are concerns that antibiotic prescribing remains suboptimal at most facilities (Table 4). Consequently STG compliance was seen as suboptimal, although relatively similar across health facilities.

Overall, all the WHO/INRUD indicators had a poor to fail rating with the AUC of the ROC (Figure 1). In addition, all the indicators showed sub-optimal sensitivity and specificity. This raises doubts about the appropriateness of the current WHO/ INRUD criteria to assess the quality of prescribing in countries with high rates of both infectious and non-infectious diseases such as Namibia and other sub-Sahara African countries. This is because our findings suggest a poor performance or accuracy of the WHO/ INRUD indicators in evaluating appropriate medicine use in PHCs. A multivariate logistic regression showed that only the antibiotic prescribing indicator independently predicted compliance to the NSTG (p < 0.001) (Table 3). These findings, together with the recent findings from PHCs in Botswana [36], are a major concern as the WHO/INRUD indicators are still widely used in Namibia and globally to monitor and report on medicine use.

Consequently, there is a need to review the use of WHO/INRUD indicators in differing levels of care, patient populations, and health sector disease states, and update these building on OMCI, PACT and other projects. This means developing new indicators that more accurately assess the quality of prescribing with their validity and reliability robustly tested using agreed methodologies [40, 41]. Possible indicators surrounding antibiotic use could include target ratios for broad to narrow-spectrum penicillins, cephalosporins and macrolides, as well as target percentage goals for combination penicillins vs. amoxycillin and for the fluorquinolones, as these have all been identified as areas of concern to increase AMR rates and adverse drug reactions [28,29, 42,43]. There could also be target BP rates for patients

with cardiovascular disease given current high prevalence rates in Namibia and concerns with adherence [44], as well as HbA1c levels in patients with type 2 diabetes, similar to initiatives in the UK [45, 46]. Prescribing targets could also be established for INN prescribing to further improve rates, with greater education around INN prescribing starting in medical school to again mirror activities in the UK with its high voluntary INN prescribing rate [46]. Potential next steps will include organizing meetings with key stakeholder groups to develop and refine potential indicators using robust methodologies to better assess the quality of prescribing in the future among PHCs in Namibia.

The main limitation of the study was that it was carried out in only one region of Namibia and with a limited number of health facilities. In addition the study was of a cross sectional design carried over a six month period, which have their own limitations. However, we believe our findings are robust based on the chosen site and their representational characteristics providing future guidance to improve medicine use in Namibia and wider.

Conculsion.

All the four WHO/INRUD indicators had low validity in predicting the quality of prescribing as outlined in the NSTG. In addition, prescribing according to the WHO indicators at health facilities in Namibia were sub-optimal. This needs to be addressed. Any developed indicators have to meaningfully assess the quality of prescribing in Namibia since only the antibiotic prescribing indicator among the WHO/INRUD indicators was an independent covariate for assessing the appropriateness of prescribing based on the NSTG. These are projects for the future.

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There was no external funding for this research. The authors have no conflict of interest to declare. The authors would like to acknowledge prescribers and pharmacists at the study sites for facilitating the data collection.

Authors contributions

NQ, HRK and DK designed the study and collected data. NQ, DK, BG, SC, and AB analysed the data. NQ and DK were responsible for the data validation. NQ and DK wrote the initial draft. BG, AM, SC, AB, HRK and DK refined successive drafts. All the authors approved the final manuscript. BG is the guarantor.

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