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ORIGINAL RESEARCH

Antibiotic Use and Resistance Among Prescribers: Current Status of Knowledge, Attitude, and Practice in Egypt

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Introduction: The problem of antimicrobial resistance (AMR) is rising worldwide. One of the most significant factors influencing antimicrobial resistance in low- and middle-income countries is the lack of the skills and knowledge of health care providers for proper antimicrobial use.

prescribers towards antibiotic use.

prescribers.

Results: Five hundred Egyptian physicians responded to the questionnaire, 113 (22.6%) reported having Antibiotic Stewardship Program (ASP) at their workplace, 99 (19.8%) use international antibiotic guidelines as the main source for prescribing activity, 358 (71.6%) recorded adequate knowledge. Elder prescribers and those who work in private or university hospitals were significantly more knowledgeable (p=0.031 and 0.001, respectively). Forty-seven (9.4%) showed a positive attitude towards proper antibiotic prescription. The type of work institution; primary health clinics and private, or university hospital and the specialty are significantly associated with a positive attitude (p=0.009 and 0.015, respectively). Seventy-eight (15.6%) expressed a proper antibiotic prescription practice. Elder age, more experienced and trained prescribers showed more proper practice. The implementation of ASP and using proper sources of information were significantly associated with proper practice (p=0.012 and 0.008, respectively).

Conclusion: Egyptian prescribers have a good level of knowledge about antibiotics. However, low rates of positive attitude and proper practice towards the problem of AMR and ASP are recorded.

Keywords: antibiotics, resistance, stewardship, prescribers, multidrug resistance, low- and middle-income countries

Introduction

Antimicrobial drugs have been widely used in human medicine for more than 50 years either as prophylaxis or therapeutics, with tremendous benefits to human health. Unfortunately, widespread usage, misuse or inappropriate prescription have resulted in the emergence of drug-resistant microbes.¹ According to World Health Organization (WHO) reports, more than 50% of all antibiotics are prescribed, sold or dispensed incorrectly. In addition, 50% of patients do not take antibiotics properly.² Infections with drug-resistant bacteria have increased not only morbidity and mortality, but also length of hospitalization and care costs.³

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Aim: To identify knowledge, characterize practices and describe the attitude of Egyptian Methods: A self-administered validated questionnaire was distributed among Egyptian

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Antibiotic use in medicine varies not only by medical condition, but also by geographical area associated with various health systems, culture and education, systemic elements of the primary care system and consumer characteristics.⁴ Field studies conducted in resourcepoor countries have highlighted a large degree of excessive use of drugs both in health facilities and in the community. This has gone a long way to add to the everincreasing cost of providing health care, particularly in the event of drug resistance.⁵ In order to ensure rational prescription of medications, prescribers must adopt the guidelines for treatment and must also follow the standard prescribing protocol.⁶ One of the most significant factors influencing antimicrobial resistance in low- and middle-income countries is the lack of the skills and knowledge required by health care providers for proper antimicrobial use.⁷

Few scattered studies have been conducted in Egypt on antibiotic use among physicians in a single facility^{8,9} and undergraduate medical students.¹⁰ It is therefore important to resolve the issues of antibiotic usage among a wide range of prescribers to explore their possible position in the battle against irrational antibiotic use. To the best of our knowledge, this is the first national study that identifies knowledge, characterizes practices and describes the attitude of Egyptian prescribers towards antibiotic use.

Materials and Methods Study Setting and Design

This study was carried out in Egypt, a country linking northeast Africa with the western part of Asia. Egypt is divided into 27 governorates and each has a capital and at least one city. Most governorates have a population density of more than one thousand per one kilometer. This cross-sectional, descriptive study of a representative sample of Egyptian Health Care Workers (HCWs) was conducted according to the international guidelines of Strengthening the Reporting for Observational Studies in Epidemiology, STROBE. The sample size required for the study was estimated to be 483, with a power of 80% and a confidence level of 95%. The study followed the principles of the Helsinki Declaration and ethical approval was obtained from the institutional review board of Zagazig University, Egypt. The completion of the questionnaire by participants was considered an agreement for participation in the study.

Data Collection Tool

A self-administered questionnaire was directed to health care personnel who prescribed antibiotics from 1st February 2020 to 30th April 2020. To develop the questionnaire, an in-depth literature review was carried out.^{3,6,8–10} An initial version of the questionnaire was drafted and then revised by the research team. The questionnaire had been piloted among a group of study participants to state all the amendments about language and simplicity of use, in addition to assess the practicability of the questionnaire. Their comments and suggestions were addressed accordingly. Data gathered from the pilot study were not included in final analysis.

This standardized questionnaire was divided into 4 sections. Section (1) included five items to provide information about the demographic and professional related data. Section (2) included four items to provide information about respondents' previous training concerning antibiotics stewardship, their source of information, the presence of Antibiotic Stewardship Program (ASP) and the application of WHO AWaRe classification of antibiotics at their workplaces "Access, Watch, Reserve groups". The third section included fourteen items to assess the participants' general knowledge on antibiotic resistance and use. Each question was answered by either yes, no, or do not know. Yes, was the correct answer to some questions and no for others. Section (4) included seventeen items to determine the respondents' attitude towards antibiotic resistance and prescription practices. Responses for the attitude section were recorded on the 5-point Likert scale as follows: strongly agree, agree, neutral, disagree, and strongly disagree. The fifth section included four items to identify the participants' practice in applying proper antibiotic prescription.

Data Collection Process

The questionnaire was distributed via different on-line applications, e.g., WhatsApp, Facebook and Messenger. A printed form was available and distributed. The title, aim and objectives of the study were clarified in the questionnaire form. Respondents were asked if they want to participate or not. A clear statement that their completion of the questionnaire implies their consent to participate was included in the form. We clarified that this survey was for research purpose. Data were collected anonymously.

Statistical Analysis

All data were collected, tabulated, and statistically analyzed using SPSS version 19. Categorical qualitative variables were expressed as absolute frequencies (number) and relative frequencies (percentage). Categorical data were compared using Chi-square test (χ 2test). All tests were two sided. p-value <0.05 was considered statistically significant (S), p-value <0.001 was considered highly statistically significant (HS), and p-value ≥ 0.05 was considered statistically insignificant (NS). For assessing knowledge, a score of one point was given if the right answer was chosen and a score of zero was given if the wrong answer or 'do not know' was chosen. A total score ranged from "0" to "28". Participants were grouped into 2 categories according to their level of knowledge: adequate (≥60% of total marks), and inadequate (<60% of total marks). For assessing attitude, a score of 5 to 1 was given to each item according to the selected point on Likert scale. Participants were grouped into positive or negative; $\geq 60\%$ or < 60% of total marks, respectively. This giving a total score ranging from 17 to 85. Attitude level $\geq 60\%$ is considered as a positive attitude. For assessing practice, antibiotic prescription practice was calculated by giving one point for each selected step. This gave a score ranging from 1 to 4. Practice level $\geq 60\%$ is considered as proper practice.¹¹

Results

A total of 500 Egyptian physicians were included in the study: 295 (59%) males and 205 (41%) females, with an age range from 25 to more than 50 years. Prescribers were distributed across the different types of Egyptian health-care institutions, the highest percentage 222 (44.4%) worked in Ministry of Health and population (MOHP) Hospitals, Table 1.

Out of the 500 respondents, 113 (22.6%) reported they have ASP at their workplace and 67 (13.4%) have lists of antibiotics classified into "Access, Watch, Reserve "groups.

The main sources of information about antibiotic resistance and ASP are international antibiotic guidelines 99 (19.8%), advice from older colleagues 85 (17%) and pharmaceutical companies 83 (16.6%), Figure 1.

Adequate knowledge is recorded for 358 (71.6%) of the participants (Table 2). A statistically significant association was observed between adequate knowledge level and age (X2=19.9, p=0.031) and work institutions (X2=19.9, p=0.001), where elder prescribers and those

who work in private or university hospitals were significantly more knowledgeable.

Forty-seven prescribers (9.4%) showed a positive attitude towards proper antibiotic prescription, Table 3. The majority 434 (86.8%) agreed on the importance to identify the resistance rate at their workplace and 404 (80.8%) expressed their needs to attend antibiotic training courses.

Most of them (70–80%) agreed that frequent antibiotic prescription; mainly broad-spectrum antibiotics for the same patient, inappropriate doses, patient demand and poor infection control practices have a role in increase the burden of antimicrobial resistance. On the other hand, the majority 406 (81.2%) consider that unindicated prescription of antibiotics does not cause harm. The type of work institution (X2=15.33, p=0.009) and the specialty (X2=10.52, p=0.015) are significantly associated with a positive attitude, where the prescribers who work in primary health clinics, private Clinic or university showed a more positive attitude.

The antibiotic prescription rates among study participants were: 1–2 times per week for 65 (13%), 3–5 times per week for 136 (27.2%), once daily or more for 277 (55.4%), once monthly for 8 (1.6%), and once weekly for 14 (2.8%) of them. Seventy-eight (15.6%) participants expressed a proper antibiotic prescription practice. A high rate of prescribers 345 (69%) order culture only when no improvement of the clinical condition is showed, 180 (36%) consult the lab when there is no clinical response with the treatment regime. Practicing different prescription steps is presented in Figure 2.

The age (X2=13.86, p=0.001), experience years (X2=11.65, p=0.009) and training (X2=9.46, p=0.002) were presented as significant factors that affect the practice. Elder age, more experienced and trained prescribers showed more proper practice. The implementation of antimicrobial stewardship program (X2=9.69, p=0.008) and using proper sources of information (X2=6.33, p=0.012) were significantly associated with proper practice. A significant positive correlation between knowledge and attitude is present and also between knowledge and practice (p < 0.005).

Discussion

The potential dangers associated with inappropriate use of antibiotics are the emergence and dissemination of resistant micro-organisms, and significant adverse effects such as increase in the duration of hospitalization, increase in the hazard of drug toxicity, and a significant increment in

Table I Egyptian Antibiotics Prescribers' Demographic Characteristics, Professional and Institutional Information

Items		Specialty				
		Medical n=313	Surgery n=141	ER and ICU n=42	Others n=4	
Gendre	Female	149 (47.6%)	42 (29.8%)	12 (28.6%)	2 (50.0%)	
	Male	164 (52.4%)	99 (70.2%)	30 (71.4%)	2 (50.0%)	
Age (Years)	25–35	183	85	28	4	
		(58.5%)	(60.3%)	(66.7%)	(100.0%)	
	36–50	101	43	13	0	
		(32.3%)	(30.5%)	(31.0%)	-	
	More than 50	29	13	1	0	
		(9.3%)	(9.2%)	(2.4%)	-	
Work Environment	Health Insurance Organization	35 (11.2%)	21 (14.9%)	7 (16.7%)	0	
	MOH Hospital	146 (46.6%)	64 (45.4%)	12 (28.6%)	-	
	Primary Health Clinic	42 (13.4%)	12 (8.5%)	0	3 (75.0%)	
	Private Clinic	8 (2.6%)	4 (2.8%)	0	l (25.0%)	
	Private Hospital	9	4	5	0	
		(2.9%)	(2.8%)	(11.9%)	-	
	University Hospital	73	36	18	0	
		(23.3%)	(25.5%)	(42.9%)	-	
Educational Level	Bachelor	84	36	14	2	
		(26.8%)	(25.5%)	(33.3%)	(50.0%)	
	Diploma	51	20	I	I	
		(16.3%)	(14.2%)	(2.4%)	(25.0%)	
	Doctorate	33	20	11	0	
		(10.5%)	(14.2%)	(26.2%)	-	
	Fellowship	16	I	1	0	
		(5.1%)	(0.7%)	(2.4%)		
	Master	129	64	15	I	
		(41.2%)	(45.4%)	(35.7%)	(25.0%)	

(Continued)

Table I (Continued).

Items		Specialty	Specialty				
		Medical n=313	Surgery n=141	ER and ICU n=42	Others n=4		
Work Experience	I–<5 years	81	46	15	2		
		(25.9%)	(32.6%)	(35.7%)	(50.0%)		
	5–10 years	101	42	10	I		
		(32.3%)	(29.8%)	(23.8%)	(25.0%)		
	Less than I year	16	2	3	0		
		(5.1%)	(1.4%)	(7.1%)	-		
	More than 10 years	115	51	14	I		
		(36.7%)	(36.2%)	(33.3%)	(25.0%)		
Did you attend any training/	No	191	112	27	2		
educational course for antibiotics prescription		(61.0%)	(79.4%)	(64.3%)	(50.0%)		
antibiotics prescription	Yes	122	29	15	2		
		(39.0%)	(20.6%)	(35.7%)	(50.0%)		
Your healthcare facility has	I do not Know	114	77	21	4		
an implemented antimicrobial stewardship		(36.4%)	(54.6%)	(50.0%)	(100.0%)		
program	No	112	46	13	0		
		(35.8%)	(32.6%)	(31.0%)	-		
	Yes	87	18	8	0		
		(27.8%)	(12.8%)	(19.0%)	-		
Do you have lists of	l do not know	128	71	16	2		
antibiotics classified into "Access, Watch, reserve " groups in your workplace		(40.9%)	(50.4%)	(38.1%)	(50.0%)		
	No	138	58	19	1		
		(44.1%)	(41.1%)	(45.2%)	(25.0%)		
	Yes	47	12	7	I		
		(15.0%)	(8.5%)	(16.7%)	(25.0%)		

Abbreviations: ER; Emergency Room. ICU; Intensive Care Unit. MOH; Ministry of Health.

the cost.³ Prescribers may modify their practice only when their attitudes, skills, beliefs and knowledge are integrated with each other potentially resulting in a decrease in antibiotic prescribing which may in turn help reduce the development and spread of infections that are resistant to antibiotics^{12,13} Thus, this study was designed to evaluate knowledge, attitude and practices of Egyptian prescribers.

In the current report, the study population is considered to be representative covering a wide spectrum of the antibiotic prescribers working in different Egyptian healthcare institutes, with different educational and experience standards. Out of the 500 respondents, (22.6%) reported they have implemented ASP at their workplace and (13.4%) have lists of antibiotics classified into "Access, Watch, Reserve" groups. These low rates underscore the need of updated national data about the challenges and barriers in implementing ASP in Egyptian healthcare settings (e.g., shortage of competent microbiology labs, poor knowledge, resistance to change among HCWs, lack of training, lack of financial support).^{14,15} Proper review of exact causes and allocation of resources to increase the implementation coverage could take place.

Information source

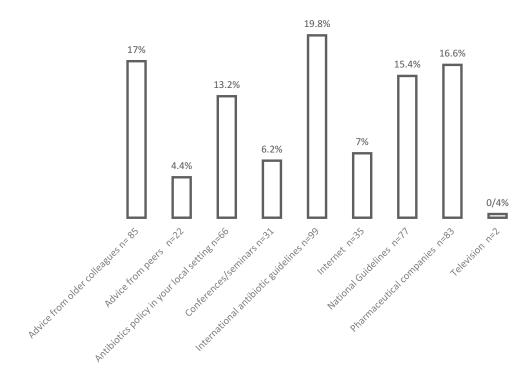


Figure I The main source of information about antibiotic resistance and stewardship.

In low- and middle-income countries, one of the most important factors affecting antimicrobial resistance is the lack of proper training/education of healthcare providers regarding proper antimicrobial use.² Similarly, in Europe, most final-year medical students lacked self-reported preparedness in antibiotic prescribing and desired further education¹⁶ and a wide variation in the exposure of students in different European countries to important principles of appropriate antibiotic prescribing was observed.¹⁷ The current report revealed that (66.4%) of the participants did not get any training/educational courses about antibiotic prescription, although most of them have completed their postgraduate studies (14.6% Diploma; 41.8% Master and 12.8% Doctorate). This finding raises concerns about the content of current undergraduate, postgraduate and continuous medical educational programs in Egypt and gives clue for the curriculum designers and the medical training providers to consider the inclusion of this vital topic in their courses. Elective courses covering topics like antibiotic prescription and antibiotic stewardship should be designed, delivered by experts and made available for both undergraduate as well as postgraduate candidates.

The presence of an accurate updated source of data about the proper use of antibiotics is crucial. Despite the availability of both national and international guidelines that offer balanced and evidence-based advice for the prescribers, only (19.8%) of the participants referred to international antibiotic guidelines as the main source of information about antibiotic resistance and stewardship program versus (15.4%) who used the national guidelines, and (13.2%) who used the antibiotics policy in their healthcare setting. Discrepant results were reported from previously published studies; a study from Ireland showed the hospital antibiotic prescribing guidelines as minor source and the dominant influence was the advice of senior doctors.¹⁸ Krishnakumar and Tsopra, 2019 stated the clinical practice guidelines as the most frequently used resource, followed by antibiotic websites, personal memos, and scientific publications.¹⁹ By default, the international guidelines should be revised by national and local authorities and adapted to the local clinical needs. Furthermore, the guidelines should proactively address prescribers' concerns.²⁰ Further insights are needed to explore the reason for this unacceptable low rate of using the national guidelines and the local antibiotics policy.

Item (Correct Answer)	Yes No (%)	No No (%)	l Do Not Know No (%)
Antibiotics kill/inhibit pathogenic and commensal bacteria (Yes)	481 (96.2)	16 (3.2)	3 (0.6)
Antibiotics kill/inhibit viruses (No)	20 (4)	474 (94.8)	6 (1.2)
All cases with coughs, colds and sore throats get improved with the use of antibiotics (No)	64 (12.8)	411 (82.2)	25 (5)
Taking antibiotics often has side-effects such as diarrhea (Yes)	403 (80.6)	73 (14.6)	24 (4.8)
In Egypt, antibiotics are available for over counter use in pharmacies (Yes)	394 (78.8)	61 (12.2)	45 (9)
The efficacy of antibiotic is higher if it is newer or of higher price (No)	106 (21.2)	322 (64.4)	40 (8)
The longer the duration of antibiotic use, the better the response (No)	115 (23)	349 (69.8)	36 (7.2)
Overuse of antibiotics makes antibiotics become ineffective (Yes)	461 (92.2)	28 (5.6)	11 (2.2)
Bacteria can become resistant to antibiotics (Yes)	476 (95.2)	15 (3)	9 (1.8)
Healthy people can carry antibiotics resistant bacteria (Yes)	451 (90.2)	18 (3.6)	31 (6.2)
Antibiotics resistance is a worldwide problem: (Yes)	435 (87)	28 (5.6)	37 (7.4)
Antibiotics resistance is a problem in Egypt: (yes)	261 (52.2)	149 (29.8)	90 (18)
Antibiotics resistant bacteria spread easily from person to person: (Yes)	241 (48.2)	81 (16.2)	178 (35.6)
New antibiotics are rapidly solve the problem of resistance: (No)	253 (50.6)	25 (5)	222 (44.4)

Table 2 Knowledge of Antibiotic Prescribers About Antimicrobial Resistance and Use

Meanwhile, the pharmaceutical industry still influences the prescribers' practice by promotional interventions with about (16%) of the respondents told that pharmaceutical companies are their main source of information. Faizullah et al 2017 also found similar prescribers' response where the marketing tactics of pharmaceutical industries play an important role in increasing the prescribing and sale of certain drugs.²¹

Successful implementation of antimicrobial stewardship (AMS) programs requires buy-in from relevant clinical stakeholders. A survey of attitudes towards antimicrobial resistance may be useful to determine the level of engagement among clinical stakeholders.²²

Proper antibiotic use entails the alignment of adequate knowledge, proper practice and a positive attitude. The current report showed an adequate knowledge for (71.6%) of the participants, a positive attitude for (9.4%) of them towards proper antibiotic prescription and (15.6%) of the participants expressed a proper prescribing practice. Different patterns were published in previous studies; adequate knowledge (62.8%), proper practice (72.9%) and a positive attitude (15%).²³ In a recent single-center Egyptian study, Tahoon et al 2020 reported adequate knowledge (39.3%), a positive attitude (85.7%) and

a proper practice (31.8%) among the investigated prescribers.⁹ These findings highlight the challenge of making antimicrobial resistance a relevant local issue among clinical stakeholders who are not easily engaged in issues pertaining to antimicrobial use at the hospital. Attitude survey and behavior analysis are highly encouraged to optimize the needed actions to improve the attitude and behavior.

The gap between knowledge, practice and attitude was confirmed in many instances. Tegagn et al 2017 reported most of their study participants do not believe that antibiotics are overused (43%) and AMR is a great problem in their facility (35.5%), 74.8% of them disagree on appropriate use of antimicrobial can lead to resistance. Moreover, 31.8% of HCP believe new antimicrobial development will keep up with the current resistance needed and 57% of them were disagreed/strongly disagreed on restriction of antibiotics for controlled use.²³

In their attitudinal survey, Cotta et al 2014 found only around one-third of respondents believed that antimicrobial resistance directly affected patients under their care.²² In a previous Egyptian study, data revealed that most healthcare providers recognize that overuse of antibiotics can lead to bacterial antibiotic resistance. Unfortunately,

Table 3 Attitude of Antibiotic Prescribers Towards Proper Antibiotic Prescription

Items	Strongly Agree/ Agree No (%)	Neutral No (%)	Disagree/Strongly Disagree No (%)
The following factors increase the antibiotic resistance rates			
• Frequent antibiotic prescription for the same patient	356 (71.2)	50 (10)	94 (18.8)
• Frequent use of broad-spectrum antibiotics for the same patient	349 (69.8)	51 (10.2)	100 (20)
• The long duration of antibiotic treatment course	297 (59.4)	89 (17.8)	114 (22.8)
 Inappropriate dosing of antibiotics 	396 (79.2)	58 (11.6)	46 (24.4)
 Misuse/abuse of antibiotics in livestock and food production 	298 (59.6)	132	70 (14)
		(26.4)	
 Patient demand for antibiotics even if it is not indicated 	386 (77.2)	28 (5.6)	86 (17.2)
 Poor infection prevention and control practices 	397 (79.4)	53 (10.6)	50 (10)
• The information delivered from pharmaceutical representatives	307 (61.4)	111	82 (16.4)
		(22.2)	
Antibiotics are abused at my workplace	333 (66.6)	55 (11)	112 (22.4)
The antibiotic is chosen according to the availability of the antibiotic more than the microbial cause	217 (43.4)	61 (12.2)	192 (38.4)
of infection			
Antibiotic prescription for patients when they do not need them, does not cause harm	42 (8.4)	25 (5)	406 (81.2)
Dispensing antibiotics without prescription should be controlled	416 (83.2)	33 (6.6)	51 (10.2)
Available antibiotics in the Egyptian market are of bad quality	192 (38.4)	131	177 (35.4)
		(26.2)	
I want to attend a training program for antibiotics prescription	404 (80.8)	38 (7)	58 (11.6)
It is important to know the resistance rates in my workplace	434 (86.8)	24 (4.8)	42 (8.4)
International guidelines are more important than local policies in antibiotic prescription	215 (43)	98 (19.6)	187 (37.4)
Some antibiotics must be ordered only by a qualified senior physician	421 (84.2)	30 (6)	49 (9.8)

because of the lack of access to microbiological testing, physicians may under-appreciate the prevalence of resistant bacterial infections.²⁴ In the United States, Giblin et al 2004 reported almost 95% of clinicians surveyed agreed that AMR was a national problem, although they were significantly less likely to perceive AMR to be a problem in their own institution or own practice.¹³ European prescribers who showed a broad awareness of AMR and its importance, with 95% agreeing AMR was a national problem. Yet fewer (63%) believed it had an impact on their own practice. They attributed the cause of AMR to pharmaceutical companies, veterinarians and poor infection control practices.²⁵

The present study stated a significant positive correlation between knowledge and attitude and a significant positive correlation between knowledge and practice. The implementation of stewardship programs and the delivery of training course significantly affect the knowledge with no statistically significant effect on the attitude. Hence, the training/educational programs should be properly designed to target the 3 learning domains: knowledge, skills and attitude. The deficiency of role models and success stories may explain this finding. A review of the design of training/education strategies, available courses, their frequency and the implementation method is highly recommended. We have to keep in mind that the time needed to change attitude is far more than that needed to give the same audience the targeted knowledge.

Limitation of Study

The specialty of physicians has an impact on their prescribing behaviors. However, in this survey, we could not analyze the features and underlying differences between all participating specialties. Further detailed studies are warranted.

Conclusion

Egyptian prescribers have a good level of knowledge about antibiotics however low levels of positive attitude and proper

Antibiotic prescritpion steps

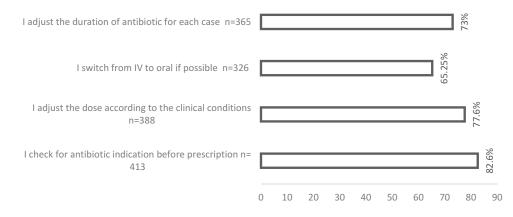


Figure 2 Prescribers practice in Antibiotic prescription.

practice towards the problem of AMR are reported. Elder, more experienced and trained prescribers showed more proper practice. The implementation of antimicrobial stewardship program and using proper sources of information were significantly associated with adequate practice. Knowledge is positively correlated with both attitude and practice.

An updated national data about challenges and barriers in implementing ASP in Egyptian healthcare settings is needed. Using regional antibiogram may be the solution for shortage of competent microbiology labs in Egyptian healthcare settings. Review of the current undergraduate, postgraduate and continuous medical educational programs in Egypt is warranted. Elective courses covering topics like antibiotic prescription and antibiotic stewardship should be designed. Attitude survey and behavior analysis are highly encouraged to optimize the needed actions to improve the attitude and behavior.

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Disclosure

The authors report no conflicts of interest in this work.

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