

International STD Research & Reviews

11(2): 37-46, 2022; Article no.ISRR.91515

ISSN: 2347-5196, NLM ID: 101666147

Knowledge and Management of Sexually Transmitted Infections by General Physicians of Lahore

Shumaila Nargus a*

^a University Institute of Public Health, University of Lahore, Pakistan.

Author's contribution

The sole author designed, analysed, interpreted and prepared the manuscript.

Article Information

DOI: 10.9734/ISRR/2022/v11i2147

Open Peer Review History:

This journal follows the Advanced Open Peer Review policy. Identity of the Reviewers, Editor(s) and additional Reviewers, peer review comments, different versions of the manuscript, comments of the editors, etc are available here:

https://www.sdiarticle5.com/review-history/91515

Original Research Article

Received 01 August 2022 Accepted 03 October 2022 Published 11 October 2022

ABSTRACT

HIV/AIDS has affected nearly every country on the planet. The effect of this surge has brought all major modes of HIV/AIDS transmission to the attention of public health professionals worldwide. Sexually Transmitted Infections (STIs), which are a major mode of transmission, increases the prevalence of HIV/AIDS. The study's aim was to assess knowledge and management of sexually transmitted infections by general Physicians of Lahore. A cross-sectional survey, of 203 general Physicians in Lahore, was conducted using structured questionnaires. Descriptive analysis was used to compute means and proportions. According to the findings of this study, 23 percent of UDS and 5 percent of GPs appropriately managed GUS syndromes. However, none of the GPs were found to managed VDS. The GPs were more likely to be trained in syndromic management who managed UDS appropriately (adj. OR: 8.0; 95% CI: 3.2-20.1), and UDS was appropriately managed (adj. OR: 2.7; 95% CI: 2.0-3.6). This study reveals a significant proportion of GPs in Karachi managed STIs. This provides a fantastic opportunity to treat and counsel STI patients and their partners. However, general Physicians lack the necessary skills. As a result, GPs must be trained in STI management in order to avert the impending disaster of HIV/AIDS. This study suggested that practicing doctors, particularly those who are young and female, should get prioritized training in STI care. Even though both the public and business sectors are actively working in this direction, additional efforts are still required.

Keywords: HIV/AIDS; sexually transmitted infections; urethral discharge syndrome; genital ulcer syndrome; vaginal discharge syndrome; general practitioner.

1. INTRODUCTION

Sexually Transmitted Infections (STIs) are spread between partners through sexual activity, most commonly vaginal intercourse, oral sex, or anal sex [1]. The infections were commonly known as venereal diseases (VD) until around 1990, when public health officials introduced the new term to improve the clarity of their public warnings. STIs probably have been around for thousands of years, but the most dangerous of these conditions, the acquired immunodeficiency syndrome (AIDS), has only been recognized in the early eighties [2]. The STI is also known as an STD (Sexually transmitted Disease).

1.1 STI Epidemiology

The STIs are caused by a host of factors categorized into; The physiological environment consist of microbiologic, hormonal and immunologic environment of the vaginal canal and /or urethra and penile lesions which determines the intra and extra-cellular presence, longevity and transmission capacity of the sexually transmitted organism The behavioural and personal environment depicts high-risk individuals' sexual behaviours and substance whereas the sociocultural environment depicts the socioeconomic, demographic, and political behaviours of the society in which the high-risk individual lives [3-4].

1.2 STI Diagnosis

STIs are currently managed in many countries using clinical or etiologic diagnosis with incomplete diagnostics [5]. This method is unreliable because many co-infections are missed.

1.3 Global Disease Burden

According to the World Health Organization (WHO), 340 million new cases of STIs are reported each year [6].

South and Southeast Asia has the highest number of new infections, estimated at 151 million, followed by Sub-Saharan Africa, Latin America, and the Caribbean. However, Sub-Saharan Africa has the highest rate of new cases per 1000 population [7].

1.4 Disease Burden at the Local Level

Pakistan does not have reliable data on the prevalence and incidence of various STIs. However, data from four provincial capitals' tertiary care centres show that 27.5 percent of 465 STI patients had gonorrhoea, 31.6 percent had syphilis, 17.2 percent had Chancroid, 18.3 percent had Herpes, and 5.2 percent had various forms of Chlamydial infections [8].

1.4.1 The HIV-STI relationship

Several studies now show that STIs, both ulcerative and non-ulcerative, increase HIV transmission [9-10]. HIV appears to change the natural history of some STIs [9]. HIV has also been isolated from both male and female genital ulcer exudates [11].

1.5 STI Health, Social, and Economic Consequences

STIs have the greatest impact on women and children [12]. Morbidity and mortality due to STIs, excluding HIV, are second only to maternal causes in women aged 15 to 44.

In Pakistan, we have all risk factors discussed in the STI epidemiology [13]. To begin with, 40.0% of Pakistanis are of reproductive age. Secondly, a study of the country's largest city found that migration accounts for 1.9 percent of urban growth (annual increase), while natural growth in urban areas accounts for 2.6 percent, annum. This pattern would be consistent with the rest of the country. Thirdly, traditional health-care providers have a huge influence on society's health-care seeking behaviours. The practice of male circumcision is most likely the most important factor responsible for slowing HIV transmission in Pakistani society.

These factors may cause Pakistan to move from a low to a high prevalence country for HIV/AIDS, implying that complacency is inappropriate.

1.6 Study's Objectives

- 1. Determine the proportion of general Physicians (GPs) who treat STI patients.
- 2. To identify the factors that influence management of STIs by GPs.

- To assess the proportion of GPs who manage the three most common STI syndromes, namely Urethral Discharge Syndrome (UDS), Genital Ulcer Syndrome (GUS), and Vaginal Discharge Syndrome (VDS) (VDS).
- To identify the factors that influence whether or not GPs in Karachi appropriately manage the three most common STI syndromes in their private practices.

2. METHODS

2.1 Research Design

The study's goals were accomplished using a descriptive cross-sectional study design. The clinics and hospitals where the GPs maintained their private practice served as the study's location. The location of this study was Lahore, Pakistan.

2.1.1 Inclusion criteria of study

- All Lahore general Physicians who see more than 40 patients per day
- 2. MBBS or a comparable degree. (As acknowledged by the Pakistan Medical and Dental Council)

2.2 Sampling Procedure

To select study units from the sample frame of 203 GPs, the simple random sampling (SRS) approach was applied. The sample size was determined using a 95% confidence level, an assumed percentage of 50% (0.5), and a 10% admissible error (0.1). The calculated sample size was 96, and when a 25% non-response rate was applied, the needed sample size was 120.

2.3 Data Gathering Instrument

To learn more about the percentage of Karachi's general Physicians that treated STIs and any possible contributing variables, a survey questionnaire in English was created. Additionally, data on the suitability of managing three STI syndromes and any relevant contributing variables was gathered.

2.4 Descriptive Analysis

The statistical package for social sciences (SPSS version 11.5 for Windows) was used to analyses the data [14]. The analysis ran the frequencies and ratios. Age, educational level,

male to female ratio, number of patients managed per day, number of STI patients managed every month, time spent with each patient, percentage of GPs handling STIs, and other factors were all given descriptive data.

2.4.1 Univariate analysis

Since the study's goals were to identify the variables that influence whether, or not a general practitioner (GP) effectively handles three STI syndromes (Objective 4), as well as whether, or not they do so, three distinct univariate analyses were carried out.

2.4.2 Multiple logistic regression

For goals 2 and 4, we created three multivariable models: one to determine whether, or not a doctor treats STI patients, and the other two to determine whether, or not a doctor treats STI syndromes, like GUS and UDS, adequately. However, as was already indicated, the model for VDS could not be created because it was discovered that none of the GP was properly managing the condition.

3. RESULTS

203 of the 332 GPs responded. This research had a response rate of 97.4%.

3.1 Percentage of GPs who Treated STI Patients

Almost all of the GPs (99.7%) managed UDS, 77.0% managed GUS, and 27.0 % managed VDS.

Percentage of doctors who treated three STI symptoms appropriately (Table 1) only 23.0% of GPs who handled UDS did it in an appropriate manner.

Table 1 shows the proportion of general Physicians (n=387) who manage three STI symptoms and the appropriateness of their knowledge of STI care. More than 387 general Physicians weree combined, indicating an overlap.

3.2 Descriptions of the Traits

3.2.1 Sociodemographic characteristics of GPs

(Table 2) Males made up 71.0% of the general Physicians (GPs) interviewed; their average age

was 40.5 years (+8.5); and they mostly completed their internships in medicine and surgery (57.9%) as opposed to medicine and gynaecology and medicine and dermatology (22.0% and 20.0% respectively).

Only a quarter of these GPs (27.1%), who make up 30.0% of the group, had ever heard of the idea of syndromic treatment of STIs.

Table 2 lists the demographics of the general Physicians (n=203) in Karachi who were treating three STI syndromes in the study. Specialised areas beyond of those already listed.

(Table 2) These doctors saw an average of 3.6 (1.2) STI patients every month, which was a small proportion of the 71.1(14.4) regular patients seen on an average day.

Table 2 lists the characteristics of Karachi's general Physicians who were

examined on management of STI patients (n=203).

3.3 Logistic Regression Analysis

Univariate analysis to determine the variables that influence whether a GP handled STI patients:

(Table 3) Compared to GPs who did not treat STI patients, those who did were more likely to manage when the number of their daily patients grew by 25, maintain patient privacy while managing STI patients, and be hired by the government.

Table 3: Univariate Odds Ratio and 95% confidence intervals (CI) of variables influencing whether or not general Physicians in Karachi, Pakistan, handled STI patients (n=203). The impact of the daily patient increase of 25 patients has been noted.

Table 1. Percentage of GPs managing three STI syndromes and appropriateness of their knowledge regarding STI management (n=387)

STI Syndrome	Number of GPs managing STI Syndromes n*	Appropriate management n (%)	Inappropriate management n (%)
Urethral Discharge Syndrome (UDS)	386	90 (23.3)	296 (76.7)
Genital Ulcer Syndrome (GUS)	298	16 (5.4)	282 (94.6)
Vaginal Discharge Syndrome (VDS)	106	0 (0.0)	106 (100)

Table 2. Practice-related characteristics of the GPs studied for managing STI patients in Karachi (n=203)

Variables	n	(%)	Mean (±SD)
Manage STI patients		, ,	
No	131	25.3	
Yes	387	74.7	
Privacy in facility			
No	121	23.4	
Yes	397	76.6	
Number of patients seen daily	203	100	71.1 (±14.4)
Number of STI patients seen per month	203	100	3.6 (±1.2)
Time spent per patient (in minutes)	203	100	2.7 (±1.3)
User fee (Rs)	203	100	83.40 (±40.30)

3.4 Multivariable Logistic Regression Model to Determine the Variables Influencing Whether a GP Consulted STI Patients or Not

(Table 4) Compared to GPs, who did not consult STI patients, STI patient-managing GPs were more likely to be under the age of 34, work for the government, and maintain their privacy.

As their daily patient load grew by 25 patients, these GPs were more likely to treat STI patients.

Table 4: Adjusted odds ratios and 95% confidence intervals (CI) of variables influencing whether, or not general Physicians in Karachi, Pakistan, consulted with STI patients (n=203).

- Based on quartile analysis, the age was dichotomized (1st quartile midpoint: 34; 2nd – 4th quartile midpoint: 34).
- 2. The impact of the average number of patients seen each day increasing by 25 patients is documented.

3.5 Univariate Analysis to Determine the Variables that Influence Whether a General Practitioner (GP) Correctly Manages Genital Ulcer Syndrome (GUS)

(Table 5) The GPs in Karachi who properly handled GUS patients were less likely to do so as their ages grew; younger GPs would have adequately controlled GUS, resulting in a fee increase of Rs. 50/-.

Table 5: Univariate Odds Ratio and 95% confidence intervals (CI) of variables related to Karachi's general Physicians' ability to treat genital ulcer syndrome effectively (n = 298). The impact of the user fee's rise of Rs. 50/- was reported.

Medicine and surgery was created by combining medicine and gynaecology and medicine.

Table 3. Multivariable Logistic Regression Model for identifying factors associated whether or not a GP managed STI patients

Variables	Manage		Odds Ratio	95% CI
	Yes	No		
Age			0.56	0.48 - 0.65
Number of patients seen daily*			3.91	2.1 - 7.2
Time spent per patient			1.46	0.97 - 2.05
(time in minutes)				
Number of STI patients seen per month			1.52	0.96 - 2.13
Privacy in facility				
No	61	59	1.0	
Yes	325	73	4.31	2.83 - 6.55
Nature of Employment				
Private	273	125	1.0	
Government	113	7	7.39	3.3 - 16.4
Training in Syndromic Management				
Never	281	97	1.00	
Ever	105	35	1.04	0.66- 1.63
Internship				
Medicine & surgery	310	102	1.00	
Medicine & dermatology	80	26	1.01	0.63-1.63

Table 4. Adjusted Odds Ratio and 95% confidence intervals (CI) of factors associated with whether or not the GPs managed STI patients in Karachi, Pakistan (n=203)

Variables	Adjusted Odds Ratio	95% CI
Age		
³ 34	1.0	
<34	18.9	8.6 – 41.7
Nature of Employment		
Private	1.0	
Government	12.2	4.5 - 33.1
Privacy in facility		
No	1.0	
Yes	2.4	1.7 – 3.4
Number of patients seen daily ²	1.34	1.13 – 1.56

3.6 Multivariable Logistic Regression Model for Determining Factors Associated With Management of Genital Ulcer Syndrome (GUS) by a General Practitioner

Table 6: Adjusted Odds Ratio and 95% Confidence Intervals (CI) of variables influencing whether general Physicians in Karachi, Pakistan, effectively handled genital ulcer syndrome (n=298).

(Table 6) Compared to the GPs who did not treat GUS correctly, the GPs who did so may have completed their internships in dermatology and medicine. The charge was raised by Rs.50/-as a result of GPs managing GUS patients properly.

The user fee's impact was stated for the Rs.50/- increase. Combining medicine and gynaecology and medicine created medicine and surgery.

Table 5. Univariate Odds Ratio and 95% confidence intervals (CI) of factors associated with whether or not the GPs appropriately managed Genital Ulcer Syndrome in Karachi (n= 298)

Variable	Manage GUS		Odds	95% CI
	Appropriate	Inappropriate	Ratio	
Age			0.90	0.83- 0.99
User fee* (in rupees)			17.8	3.8 - 83.1
Number of STI patients seen per month			1.04	0.90-1.10
Number of patients seen daily			1.2	0.97 - 1.40
Time spent per patient			1.03	0.80 - 1.25
Internship				
Medicine & Surgery ^a	15	228	1.0	
Medicine & Dermatology	35	20	26.6	12.4 - 56.8
Nature of employment				
Private	153	60	1.0	
Government	71	14	1.98	1.04-3.7
Privacy				
No	5	16	1.0	
Yes	71	206	1.10	0.39-3.09
Training in syndromic				
management				
Never	79	15	1.0	
Ever	174	30	1.1	0.56 - 2.15

Table 6. Adjusted Odds Ratio and 95% confidence intervals (CI) of factors associated with whether or not the GPs appropriately managed Genital Ulcer Syndrome in Karachi, Pakistan (n=298)

Variables	Adjusted Odds ratio	95% CI	
Internship			
Medicine & surgery ^a	1.0		
Medicine & dermatology	18.2	5.52 - 100.9	
User fee*	3.8	1.9- 7.5	

Table 7. Univariate Odds Ratio and 95% confidence intervals (CI) of factors associated with whether the GPs appropriately managed Urethral Discharge Syndrome in Karachi, Pakistan (n = 386)

Variable	Manage UDS		Odds	95% CI
	Appropriate	Inappropriate	Ratio	
Age			0.90	0.80 - 0.98
Number of STI			2.5	2.0 - 3.2
patients per month				
User fee			1.0	0.9 – 1.1
Time spent per patient			1.05	0.92 - 1.4
Number of patients seen			1.1	0.96 - 1.5
daily				
Training in Syndromic				
Management				
Never	30	251	1.0	9.4 - 27.1
Ever	69	36	16.0	
Employment				
Private	67	206	1.0	
Government	46	67	2.1	1.32- 3.32
Privacy				
No	35	26	1.0	
Yes	231	94	1.82	1.22-3.19

Table 8. Adjusted Odds Ratio and 95% confidence intervals (CI) of factors associated with whether or not the GPs appropriately managed Urethral Discharge Syndrome in Karachi (n=386)

Variables	Adjusted Odds Ratio	95% CI
Training in Syndromic		
Management		
No	1.0	
Yes	8.0	3.2 – 20.1
Number of STI patients seen	2.7	2.0 - 3.6

3.7 Univariate Study was Conducted to Determine the Variables that Influence Whether a General Practitioner (GP) Correctly Addressed Urethral Discharge Syndrome (UDS)

(Table 7) General Physicians who handled UDS well were more likely than those who did not to

have had training in syndromic treatment and to be hired by the government.

GPs were more likely to treat UDS effectively as the number of STI patients rose.

GPs were less likely to treat UDS patients properly as they aged.

Table 7: Univariate Odds Ratio and 95% confidence intervals (CI) of variables influencing whether general Physicians in Karachi, Pakistan, treated urethral discharge syndrome correctly (n = 386). Zero cell counts precluded further analysis of internship.

3.8 Multivariable Logistic Regression Model to Determine the Variables that Influence Whether or Not a GP Managed UDS

(Table 8) Compared to GPs who improperly handled UDS, the former were more likely to have received training in syndromic care. (adj.OR: 8.0; 95%CI: 3.2 - 20.1) These GPs were more likely to handle UDS well as the number of STI patients rose (adj.OR: 2.7; 95% CI: 2.0 - 3.6).

Adjusted Odds Ratio and 95% Confidence Intervals (CI) of variables influencing whether Karachi's general Physicians treated urethral discharge syndrome correctly (n = 386).

4. DISCUSSION

According to the current survey, three-fourths of Karachi's general Physicians manage STIs. However, it does not accurately depict the situation because not all of these clinicians do so. The potential to treat STI patients and their partners as well as to counsel them on many elements of risk factor reduction is greatly diminished by the poor level of proficiency in STI care. As a result, there is little overall impact on breaking the chain of STI transmission.

According to research done in Karachi, 80.0% of GPs treated STIs [15].

Even then, only 44.0% of the GPs were found to appropriately manage UDS, compared to 23.0% in the current research. According to the current study, even a lower percentage of GPs were discovered to be correctly managing GUS and none managed VDS. Because the current study adhered to the WHO standards for the care of syndromic conditions, the mismatch in percentage was discovered.

The current study indicates that there are extremely few STI patients visiting general Physicians. In this study, a general practitioner (GP) treated a maximum of twelve STI patients in a month, with a mean of 3.6 (1.2). We assume that cultural influences, [16–17] a lack of

knowledge about the person engaging in risky behaviours related to STIs [18] and inappropriate management provided by the GP are the factors that drive the patient to seek assistance from the informal sector or, more likely, be content with self-medication [16]. This amplifies the need for increased public knowledge of STIs and for GP training in STI care. The current study also reveals that GPs who treated STIs were more likely to be under 34 years of age and work for the government.

The GPs with effective GUS and UDS management were more likely to be skill-based. Studies suggest that in areas where healthcare professionals have received syndromic training, the outcomes have significantly improved [19–20]. In the current investigation, it was discovered that, none of the GPs were managing VDS) well. This critical but sometimes overlooked aspect of VDS care have to be the focus of CME programmes.

Although the incidence of HIV/AIDS is still low in Pakistan, the rise in the number of HIV/AIDS patients, over the past several years, is a strong indicator that preventative measures must be developed. Investments in STI awareness, in general, and training of GPs in identifying and treating at least major STI syndromes, in particular, are projected to significantly reduce the approaching burden of STIs and HIV/AIDS. This is because STIs play a significant role in the transmission of HIV/AIDS.

5. CONCLUSION

The results of this study show that while though three-fourths of Karachi's general Physicians (GPs) treat STI patients in their private practises, only a small percentage of these clinicians are capable of providing proper care. This study also demonstrates that the management of STIs is not reliant on knowledge or expertise. In the current study, it was discovered that GPs' understanding of VDS was insufficient, bringing attention to the problem of gender equity in access to effective STI treatment.

6. RECOMMENDATIONS

This study suggests that STI management be covered in the medical curriculum at the undergraduate level. The curriculum should be done in order to introduce the idea of syndromic management to all medical graduates. This study suggests that practising doctors, particularly

those who are young and female, should get prioritised training in STI care. Even though both the public and business sectors are actively working in this direction, additional efforts are still required.

CONSENT

The general Physicians (GPs), who later participated in the study, provided an informed consent.

ETHICAL APPROVAL

The University of Lahore's Research Ethics Committee and University Institute of Public Health Committee awarded its ethical permission.

COMPETING INTERESTS

Author has declared that no competing interests exist.

REFERENCES

- Webster's New World™ Medical Dictionary, 2nd Edition. Wiley Publishing, Inc. Available:http://www.medicinenet.com/scrip t/main/art.asp
- AIDS Primer.
 Available:http://treasuresoftheinternet.org/health/aids/us/us_aids.shtml
- Textbook of Medicine, edited by Souhami RL, Moxham J. Elsevier Health Sciences Publication Date: 1 Jan 2002. ISBN: 0443064644.
- 4. Grosskurth H, Mayaud P, Mosha F, et al. Asymptomatic gonorrhoea and chlamydial infection in rural Tanzanian men. BMJ. 1996;312:277–80.
- Ones RB, Wasserheit JN. Introduction to the biology and natural history of sexually transmitted diseases. In: Wasserheit JN, Aral SO, Holmes KK, eds. Research Issues in Human Behavior and Sexually Transmitted Diseases in the AIDS Era. Washington, DC: American Society for Microbiology. 1991;11-37.
- Wasserheit JN. Effect of changes in human ecology and behavior on patterns of sexually transmitted diseases, including human immunodeficiency virus infection. Proc Natl Acad Sci. 1994;91: 2430-435.

- 7. The Social Organization of Sexuality: Sexual Practices in the United States. Edward O Laumann, Robert T Michael, Stuart Michaels, John H Gagnon p. 377.
- 8. The world's youth: A special focus on reproductive health. Washington, DC: Population Reference Bureau and the Center for Population Options; 1994.
- 9. Brookman RR. Adolescent sexual behavior. In: Holmes KK, Mardh PA, Sparling PF, et al. Sexually transmitted diseases. New York: McGraw-Hill Inc. 1990:77-84.
- Qidwai W. Knowledge about Sexually Transmitted Infections among Young Pakistani Men. J Pak Med Assoc. 2002; 52(6): 267-8.
- Behets MT, Desormeaux J, Joseph D, et al. Control of sexually transmitted diseases in Haiti: Results and implications of a baseline study among pregnant women living in Cité, Soleil shantytowns. J Infect Dis. 1995;172:764-771.
- Over M, Piot P. HIV infection and sexually transmitted diseases. In: Jamison DT, Mosley WH, Measham AR, Babadilla JL, eds. Disease control priorities in developing countries. New York: Oxford University Press. 1993;445-529.
- Laga M. Epidemiology and control of sexually transmitted diseases in developing countries. Sex Transm Dis. 1994;21(Suppl 2):S45—S50.
- Carael, M, Cleland J, Adeokun L, et al. Overview and selected findings of sexual behaviour surveys. AIDS. 1991;5(Suppl 1): S65—S74.
- 15. Crabbe F, Carsauw H, Buve A, et al. Why do men with urethritis in Cameroon prefer to seek care in the informal health sector? Genitourin Med. 1996;72:220-2.
- Van der Geest S. Self-care and the informal sale of drugs in South Cameroon. Soc Sci Med. 1987;25:293-305.
- 17. Khamboonruang C, Beyer C, Natpratan C, et al. Human immunodeficiency virus infection and self-treatment for sexually transmitted diseases among northern Thai men. Sex Transm Dis. 1996;23:264-9.
- 18. Pallikadavath S, Sanneh A, Jenny MM, Stones RW et al. Rural women's knowledge of AIDS in the higher prevalence states of India: Reproductive health and sociocultural correlates. Health Promot. Int. 2005;20:249-259.
- Kamali A, Quigley M, Nakiyingi J, et al. Syndromic management of sexually

transmitted infections and behaviour change interventions on transmission of HIV-1 in rural Uganda: A community randomised trial. Lancet. 2003;361: 645-52.

 Mayaud P, Mabey D. Cutting edge review—managing sexually transmitted diseases in the tropics: Is a laboratory really needed? Trop Doct. 2000;30: 42-6.

© 2022 Nargus; This is an Open Access article distributed under the terms of the Creative Commons Attribution License (http://creativecommons.org/licenses/by/4.0), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Peer-review history:
The peer review history for this paper can be accessed here:
https://www.sdiarticle5.com/review-history/91515