ORIGINAL ARTICLE



Crit. Care Innov. 2021; 4(4):1-8



Symptom analysis of confirmed covid-19 patients - Cross sectional study.

Khadija Asif ¹ - A,B,C,G,H,J,K,N,O. O ORCID www.orcid.org/0000-0002-9914-2492 Farhat Abbas ¹ - E,F,I,K,L,M,O. O ORCID www.orcid.org/0000-0002-1931-2413

¹ Department of Medicine, Allied Hospital, Faisalabad Medical University (PMC), Faisalabad, Pakistan.

Address for correspondence: Khadija Asif, MD - Unit of Medicine (PMDC-97676-P) Department of Medicine, Allied Hospital, Faisalabad Medical University, Faisalabad(PMC), 01 Jail Road adjacent to Sargodha Road. 38030, Faisalabad city, Pakistan; Tel: 0092-302-330-2300; E-mail: dr.khadija.asif@outlook.com

ABSTRACT

INTRODUCTION: Coronavirus can cause respiratory disease ranging from mild upper respiratory tract illness to severe pneumonia, acute respiratory distress syndrome, and death. The purpose of this research was to analyze the relationship of gender and age with symptoms of COVID-19 disease.

MATERIAL AND METHODS: Consecutive sampling methodology was used, and all real-time polymerase chain reaction (RT-PCR) confirmed patients of COVID-19 were included. Data regarding age, gender, and symptoms with onset was recorded and analyzed using Chi-square and Student t-tests with P-value ≤ .05 considered as significant. Data was entered and analyzed using Statistical Package for Social Sciences (SPSS) version 25.0

RESULTS: Ninety-eight patients, 68 (69.4%) males and 30 (30.6%) females were included in the study. Fifty-five (56.1%) were symptomatic and the rest were asymptomatic. Mean duration of symptoms in symptomatic cases was 2.8±1.1 days. Of these, 60 (61.85%) were older than 40 years and 17 (17.34%) had a history of travel. Fever and persistent cough were reported by more than 50% patients. Fatigue, sore throat, shortness of breath and myalgias and headaches were reported by 32-21% patients. Nausea, Vomiting, runny nose, sputum, diarrhea and anorexia were reported by less than 11% patients. Fever, dyspnea and nausea were more in individuals over 40 years. Dyspnea was more in males and fatigue was more in females.

CONCLUSIONS: Many of our COVID-19 infected patients were symptomatic with fever and cough as the most common symptoms. Recognition of the pattern of these symptoms is essential for early diagnosis and management of the disease.

KEY WORDS: Coronavirus, Cough, COVID-19, Fever, Symptoms.

© 2021 The Authors. This article is an Open Access article distributed under the terms and conditions of the CC BY-NC license. https://creativecommons.org/licenses/by-nc/4.0/

ISSN 2545-2533

Received: 21.10.2021 Accepted: 29.10.2021 Published: 31.12.2021

Author Contributions (CRediT Taxonomy):

Conceptualization - **A** Data Curation - **B** Formal Analysis - **C**

- Funding Acquisition D
 - Investigation E
 - Methodology F

Project Administration - G

- Resources ${\bf H}$
- Software I
- Supervision J

Validation - K

- Visualization L
- Writing (Draft Preparation) **M** Writing (Review & Editing) - **N**
- Approved the final version **O**

OWARZYSTWO

OMOCY

DORAZNEJ

INTRODUCTION

Coronavirus belongs to the family Coronaviridae [1,2]. Coronavirus disease 2019 (COVID-19) is a respiratory disease caused by a novel coronavirus that was first reported in December, 2019 at Wuhan, China. This highly infectious disease manifests at first as dry cough, fever, myalgia and difficulty in breathing. On the other side of the spectrum, this disease may manifest as acute respiratory distress syndrome, septic shock, bleeding and coagulopathy [3,4]. Though coronaviruses are generally thought to be zoonotic, COVID-19 is believed to spread from human to human [5,6]. This virus has been spreading very quickly and has prevailed over more than 200 countries and territories causing more than 219 million cases and 4.55 million deaths worldwide until 01 October-2021. In Pakistan, till 01 October 2021, 2.16Million diagnosed cases of Covid-19 with almost 28k deaths have been reported so far [7-9]. Symptom-based evaluation of COVID-19 patients in our population is deficient. The use of laboratory investigations for diagnosis of COVID-19 infection needs to be on a large scale, keeping in mind the pandemic nature of the disease [10]. This can exhaust resources of even the developed countries let alone the developing ones. In this situation, an understanding of COVID-19 symptomatology becomes an utmost necessity for the initial screening of suspected cases and to prioritize them for further investigations. The objective of this study was to note the symptoms of Real-Time Polymerase Chain Reaction (RT-PCR) confirmed COVID-19 patients. Additionally, the age group wise differences in symptomatology were also sought.

MATERIAL AND METHODS

This cross-sectional study was conducted at Allied hospital (Faisalabad Medical University) Pakistan, In the COVID-19 patient management center of Allied hospital (Faisalabad Medical University) Pakistan, during August-2021. By consecutive sampling, 98 RT-PCR positive, confirmed cases of COVID-19 were included in the study. Patients with comorbidities such as chronic respiratory and hematological illnesses, neoplastic disorders, endocrinopathies, immunodeficiency states, gastrointestinal diseases, and cardiovascular disease were excluded. Five patients were excluded from the study due to presence of comorbidities. Asymptomatic patients were followed up regularly to observe the development of symptoms. After their discharge, they were also contacted through telephone to confirm their asymptomatic status. Age, gender, symptoms and their duration, and travel history were sought from all the patients. Data was entered and analyzed using Statistical Package for Social Sciences (SPSS) version 25.0. Quantitative variables like age and duration of symptoms were reported as mean and standard deviation. Qualitative ones like gender, symptoms, and travel history were recorded as frequencies and percentages. The presence or absence of symptoms and the individual symptoms were compared between, age groups, gender, and travel history. Individual symptoms were also compared between different age groups. Chi-square and Student t-tests were used for this purpose. *P*-value \leq .05 was considered significant.

Ethical Approval

This study was approved by the Institutional ethics committee of Allied hospital (Faisalabad Medical University) with approved no. AHF-379-FMU-04/15. Informed consent was taken from the patients.

RESULTS

Out of the 98 patients, 55 (56.1%) were symptomatic and the rest were asymptomatic. Mean duration of symptoms in symptomatic cases was 2.8±1.1 days. Median time from symptom onset to admission was 3 days. Sixty-Eight (69.4%) were male and 30 (30.6%) were females. Of these 39 (57.4%) males and 16 (53.3%) females were symptomatic, respectively. The mean patient age was 45.5±17.8 years with 38 (38.77%) were younger than 40 years and 60 (61.85%) older than 40 years. Of these 20 (52.6%) and 35 (58.3%) patients were symptomatic in each group, respectively. Seventeen (17.34%) had a history of travel with 68.8% symptomatic while 81 (83.50%) did not have a travel history with 53.1% symptomatic individuals (Table 01).

Parameter Mean age (years)		Total	Symptomatic	Asymptomatic	70 1	
		n=98 (100%)	n= 55 (56.1%)	n= 43 (43.87%)	<i>P</i> -value	
		45.5±17.8	46 ± 15.8	44.93 ± 19.46	.774	
Age Quartiles	<40	38 (38.77%)	20 (52.6%)	18 (47.4%)	2 P	
(years)	≥40	60 (61.22%)	35 (58.3%)	25 (41.7%)	.579	
	Male	68 (69.38%)	39 (57.4%)	29 (42.6%)	į.	
Gender	Female	30 (30.61%)	16 (53.3%)	14 (46.7%)	.712	
	Yes	17 (17.34%)	11 (68.8%)	6 (31.3%)	5	
History of Travel	No	81 (82.65%)	43 (53.1%)	38 (46.9%)	.249	

Table 01. Comparison of age. gender and travel history of symptomatic and asymptomatic patients.

*P-value < .05 (Significant Statistically)

OWARZYSTWO

OMOCY

DORAZNEJ

Fever and persistent cough were observed in more than 50% of the patients as shown in Figure-1. Fatigue, sore throat, shortness of breath, myalgias and headaches were reported by 32.7%, 29.1%, 27.3%, 27.3% and 21.8% of our patients. Nausea, Vomiting, runny nose, sputum, diarrhea and anorexia were reported by less than 11% patients (Figure 01).

There was a significant difference in the presence or absence of fever between age groups with only 26.3% of the under 40-year-old having fever as compared to 48.3% of over 40-year-olds (P = .03) Regarding presence or absence of dyspnea, 2.6% of under 40-year-olds had dyspnea as compared to 23.3% of over 40-year-olds (P = .006). The difference in the presence of dyspnea between genders was significant with 20.6% males reporting dyspnea compared to 3.3% females (P = .029) (Table 02). There was a significant difference between age groups in the presence or absence of nausea (P = .044), as well as presence or absence of fatigue between genders (P = .011) (Table 03).

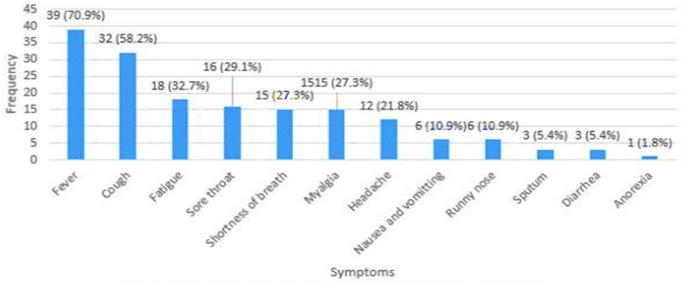


Figure 01. Symptoms of patients presenting with COVID-19

Variable		Fever n (%)	No fever n (%)	<i>P-</i> value*	Dyspnea n (%)	No Dyspnea n (%)	P-value'
Age	<40 years old	10 (26.3)	28 (73.7)	.03	1 (2.6)	37 (97.4)	.006
Quartile	≥40 years old	29 (48.3)	31 (51.7)		14 (23.3)	46 (76.7)	
Gender	Male	29 (42.6)	39 (57.4)	.385	14 (20.6)	54 (79.4)	.029
	Female	10 (33.3)	20 (66.7)		1 (3.3)	29 (96.7)	

*P-value < .05

(Significant Statistically)

Table 03. Relationship of fatigue and nausea with age groups and gender.

Variable	1 39	Fatigue present n (%)	No fatigue n (%)	<i>P</i> -value [*]	Nausea n (%)	No Nausea n (%)	<i>P</i> -value'
Age	<40 years old	6 (15.8)	32 (84.2)		0 (0)	38 (100)	
Quartile	≥40 years old	12 (20)	48 (80)	0.600	6 (10)	54 (90)	0.044
Gender	Male	8 (11.8)	60 (88.2)	0.011	3 (4.4)	65 (95.6)	0.288
	Female	10 (33.3)	20 (66.7)		3 (10)	27 (90)	

*P-value < .05

(Significant Statistically)



DISCUSSION

Clinical features of COVID-19 may vary depending on the region. They may differ even in the same country. According to a study, clinical features of COVID-19 at Wuhan were different from those in other areas of China [10]. The reason behind this might be the different host, environment and agent factors. Important findings of our study are male preponderance, mean age less than 40 years, only 60% of COVID-19 patients managed were symptomatic, fever and shortness of breath were commonest symptoms noted in more than 50% patients, and anorexia, myalgia and fatigue were relatively uncommon symptoms. Males show a comparatively weaker immune response to infectious agents as compared to females [11-13]. This is due to the difference in hormones between the sexes, the difference in the number of copies of X-linked denes related to immunity, and the difference in the disease susceptibility genes between males and females [11,12]. This and possibly more outdoor exposure may increase the chances of male gender predominance with reference to COVID-19 infection as has been noted in various studies [14,15]. In a Chinese COVID-19 analysis focusing confirmed patients, male to female ratio was 54.4%:48.6% [16]. We also noted male predominance in our study with 69.4% of the male patients. Around 40% of our patients were less than 40 years old. According to an analysis, 27.2% of COVID-19 patients managed in China were younger than 40 years and 86.6% patients were in the age range of 30-79 years [16]. In the study by Huang et al., the median age was 49 years while in the study by Xu et al., it was 41 years [10,14]. The reason for the lesser median age in our patients is that the Pakistani population mainly consists of younger persons with a median age of 22.8 years [17].

Travel history of countries where COVID-19 infection is prevalent is important in the context of the spread of infection. According to a study, travel history is more important than chest radiograph findings for early detection and isolation [18]. The first case in the USA had also traveled from China [19]. About a quarter of our patients had a positive travel history while the rest had locally transmitted infections. Interestingly none had visited China. Persons with RT-PCR confirmed COVID-19 infection who neither had any symptoms at the time of test nor did they showed any symptoms on follow-up were termed as asymptomatic laboratory confirmed cases. About 5%-80% of persons who are mostly children and young adults are tested positive, but do not have symptoms according to an analysis [20]. Asymptomatic cases are important in the spread of the disease as a false sense of security might compromise the practice of preventive measures. Asymptomatic patients can also be a source of continuous infection and a sustained level of infected cases in the community. They can become a hurdle in complete eradication of the disease even if the pandemic is over. According to WHO, although asymptomatic transmission of COVID-19 is rare according to the available data, but it cannot be ignored [21]. A study reports the transmission of COVID-19 from one asymptomatic family member to five members of the same family [22]. More than forty percent of our COVID-19 confirmed cases were asymptomatic. Out of these, 47.4% were less than 40 years old. A better immune response can be the reason for the same in young infected persons [23]. In a study by Xu et al., cough (81%) and fever (77%) were the commonest symptoms [10]. In another study by Huang et al., fever was present in 98%, cough in 76%, and sputum production in 28.6% of the patients [14]. About 1.4-7.6% of mild and 32.6% of



severe COVID-19 infected patients complained of shortness of breath [24]. According to Menni et al., anosmia was present in 59.4% of COVID-19 cases.[25] Out of the 56% symptomatic cases in our study, fever (61.9%) and cough (57%) were most frequently noted. We did not observe anosmia in any patient. In our study, fatigue, anorexia, and myalgia were least reported symptoms, although these have been reported in 44-50% of COVID-19 patients [10,14]. Interestingly, anorexia and myalgia were significantly more common in our older patients. Time from symptom onset to admission in our patients was three days. This is lesser than the median time of 6.5 days reported in a previous study [10]. This might be due to the anxiety of the community towards minor symptoms of the disease, when mild to moderate disease can be managed safely at home.

This study focused on laboratory confirmed COVID-19 patients as RIU&T is the center point of such patient management of the Allied hospital (Faisalabad Medical University) Pakistan. The majority of these patients were diagnosed with contact tracing rather than attending a health care facility for symptoms. This along with the fact that many infected persons are asymptomatic may have skewed our findings. Catheter, it will help to give lavage directly in distal lung subsegments as in blind-mini-BAL and obtain uncontaminated samples without major disconnections and direct contact with secretions.

CONCLUSIONS

Many of our COVID-19 infected patients were symptomatic. Fever and cough were the commonest symptoms in these patients. Fatigue, Sore throat, shortness of breath, myalgias and headaches were also reported by some. Nausea, vomiting, runny nose, sputum, diarrhea and anorexia were the least reported symptoms. Individuals older than 40 years are more likely to be symptomatic regardless of gender. Recognition of the pattern of these symptoms is essential for early diagnosis and management of the disease.

SUPPLEMENTARY INFORMATION

Funding: This research received no external funding. **Institutional Review Statement:** The study was conducted according to the guidelines of the Declaration of Helsinki. **Informed Consent Statement:** Informed consent was obtained from all subjects involved in the study. **Data Availability Statement:** The datasets generated and analyzed during the current study are available from the corresponding author on reasonable request. **Conflicts of Interest:** The authors declare no conflicts of interest.



REFERENCES

- [1] Fehr AR, Perlman S. Coronaviruses: an overview of their replication and pathogenesis. Methods Mol Biol. 2015; 1282: 1-23.
 doi: https://doi.org/10.1007/978-1-4939-2438-7
- [2] Q&A on coronaviruses (COVID-19). WHO.
 [WWW]: https://www.who.int/news-room/q-a-detail/q-a-coronaviruses (accessed 27 March 2020)
- [3] Wu Ch, Chen X, Cai Y. Risk Factors Associated With Acute Respiratory Distress Syndrome and Death in Patients With Coronavirus Disease 2019 Pneumonia in Wuhan, China. JAMA Intern Med. 2020; 180(7): 934-943. doi: https://doi.org/10.1001/jamainternmed.2020.0994
- [4] Chen N, Zhou M, Dong X, Qu J, Gong F, Han Y, et al. Epidemiological and clinical characteristics of 99 cases of 2019 novel coronavirus pneumonia in Wuhan, China: a descriptive study. Lancet 2020; 395(10223): 507-513.
 doi: https://doi.org/10.1016/S0140-6736(20)30211-7
- [5] Chan JF-W, Yuan S, Kok KH, To KK, Chu H, Yang J, et al. A familial cluster of pneumonia associated with the 2019 novel coronavirus indicating person-to-person transmission: a study of a family cluster. Lancet 2020; 395(10223): 514-523. doi: https://doi.org/10.1016/S0140-6736(20)30154-9
- [6] Zhou P, Yang XL, Wang XG, Hu B, Zhang L, Zhang W, et al. A pneumonia outbreak associated with a new coronavirus of probable bat origin. Nature 2020; 579(7798): 270-273. doi: https://doi.org/10.1038/s41586-020-2012-7
- [7] Coronaviruse, situation reports. WHO 2020.
 [WWW]: https://www.who.int/docs/default-source/coronaviruse/situationreports/20200415-sitrep-86-covid-19.pdf?sfvrsn=c615ea20_2 (accessed 15 April 2020)
- [8] Covid.gov.pk. 2020. [WWW]: http://covid.gov.pk/ (accessed 8 April 2020)
- [9] Naming the coronavirus disease (COVID-19) and the virus that causes it. WHO 2020. [WWW]: https://www.who.int/emergencies/diseases/%20novel-coronavirus2019/technical%20guidance (accessed 8 April 2020)
- [10] Xu XW, Wu XX, Jiang XG, Xu KJ, Ying LJ, Ma CL, et al. Clinical findings in a group of patients infected with the 2019 novel coronavirus (SARS-Cov-2) outside of Wuhan, China: retrospective case. BMJ. 2020; 368: m606. doi: https://doi.org/10.1136/bmj.m606
- [11] Klein SL, Flanagan KL. Sex differences in immune responses. Nature Rev Immunol. 2016; 16(10): 626–638. doi: https://doi.org/10.1038/nri.2016.90
- [12] Rettew JA, Huet-Hudson YM, Marriott I. Testosterone reduces macrophage expression in the mouse of tolllike receptor 4, a trigger for inflammation and innate immunity. Biol Reprod. 2008; 78(3): 432–437. doi: https://doi.org/10.1095/biolreprod.107.063545
- [13] Bouman A, Heineman MJ, Faas MM. Sex hormones and the immune response in humans. Hum Reprod Update. 2005; 11(4): 411–423. doi: https://doi.org/10.1093/humupd/dmi008
- [14] Huang C, Wang Y, Li X, Ren L, Zhao J, Hu Y, et al. Clinical features of patients infected with 2019 novel coronavirus in Wuhan, China. Lancet. 2020; 395(10223): 497–506. doi: https://doi.org/10.1016/S0140-6736(20)30183-5



- [15] MERS Clinical Features. Centers for Disease Control and Prevention. 2020. [WWW]: https://www.cdc.gov/coronavirus/mers/clinical-features.html (accessed 8 April 2020)
- [16] Epidemiology Working Group for NCIP Epidemic Response, Chinese Center for Disease Control and Prevention. The epidemiological characteristics of an outbreak of 2019 novel coronavirus diseases (COVID-19) in China. Zhonghua Liu Xing Bing Xue Za Zhi . 2020; 41(2): 145-151. doi: https://doi.org/10.3760/cma.j.issn.0254-6450.2020.02.0035
- [17] Robinson DP, Huber SA, Moussawi M, Roberts B, Teuscher C, Watkins R, et al. Sex chromosome complement contributes to sex differences in coxsackievirus B3 but not influenza A virus pathogenesis. Biol Sex Differ. 2011; 2: 8. doi: https://doi.org/10.1186/2042-6410-2-8
- [18] Pakistan Population Worldometer: Worldometers. 2020. [WWW]: https://www.worldometers.info/world-population/pakistan-population/ (accessed 8 April 2020)
- [19] Holshue ML, DeBolt C, Lindquist S, Lofy KH, Wiesman J, Bruce H, et al. First case of 2019 novel coronavirus in the United States. N Engl J Med. 2020; 382(10): 929-936. doi: https://doi.org/10.1056/NEJMoa2001191
- [20] COVID-19: What proportions are asymptomatic? CEBM. 2020. [WWW]: https://www.cebm.net/covid-19/covid-19-what%20-proportion-are-asymptomatic/ (accessed 8 April 2020)
- [21] Coronaviruse, situation reports. WHO 2020. [WWW]: https://www.who.int/docs/%20default-source/coronaviruses/situation-reports%20/20200402-sitrep-73covid-19.pdf?%20sfvrsn=5ae25bc7_2 (accessed 8 April 2020)
- [22] Bai Y, Yao L, Wei T, Tian F, Jin D, Chen L, et al. Presumed Asymptomatic Carrier Transmission of COVID-19. JAMA. 2020; 323(14): 1406-1407. doi: https://doi.org/10.1001/jama.2020.2565
- [23] Simon AK, Hollander GA, McMichael A. Evolution of the immune system in humans from infancy to old age. Proc Biol Sci. 2015; 282(1821): 20143085. doi: https://doi.org/10.1098/rspb.2014.3085
- [24] Young BE, Ong SWX, Kalimuddin S, Low JG, Tan SW, Loh J, et al. Epidemiologic Features and Clinical Course of Patients Infected with SARS CoV-2 in Singapore. JAMA. 2020; 323(15):1488-1494. doi: https://doi.org/10.1001/jama.2020.3204
- [25] Menni C, Valdes AM, Freidin MB, Ganesh S, Moustafa JS, Visconti A, et al. Loss of smell and taste in combination with other symptoms is a strong predictor of COVID-19 infection. Nat Med. 2020; 26: 1037–1040. doi: https://doi.org/10.1038/s41591-020-0916-2