

Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.

FISEVIER

Contents lists available at ScienceDirect

Journal of Infection and Chemotherapy

journal homepage: http://www.elsevier.com/locate/jic



Original Article

Clinical characteristics of adult patients hospitalized with laboratory-confirmed COVID-19 pneumonia



Elif Sargin Altunok ^{a, *}, Mustafa Alkan ^a, Sadettin Kamat ^b, Berna Demirok ^a, Celal Satici ^b, Mustafa Asim Demirkol ^b, Bengul Gursoy ^b, Cemile Dilsah Surmeli ^b, Ferhat Cengel ^c, Mustafa Calik ^d, Ulku Aygen Turkmen ^e

- ^a Infectious Diseases and Clinical Microbiology, Istanbul Gaziosmanpasa Training and Research Hospital, Istanbul, Turkey
- ^b Chest Diseases, Istanbul Gaziosmanpasa Training and Research Hospital, Istanbul, Turkey
- ^c Radiology, Istanbul Gaziosmanpasa Training and Research Hospital, Istanbul, Turkey
- ^d Emergency Medicine, Istanbul Gaziosmanpasa Training and Research Hospital, Istanbul, Turkey
- ^e Anesthesia and Reanimation, Istanbul Gaziosmanpasa Training and Research Hospital, Istanbul, Turkey

ARTICLE INFO

Article history: Received 18 June 2020 Received in revised form 15 September 2020 Accepted 15 October 2020 Available online 23 October 2020

Keywords: COVID-19 Coronavirus Clinical characteristics Outbreaks Pneumonia Turkey

ABSTRACT

Background: The clinical spectrum of COVID-19 has a great variation from asymptomatic infection to acute respiratory distress syndrome and eventually death. The mortality rates vary across the countries probably due to the heterogeneity in study characteristics and patient cohorts as well as treatment strategies. Therefore, we aimed to summarize the clinical characteristics and outcomes of adult patients hospitalized with laboratory-confirmed COVID-19 pneumonia in Istanbul, Turkey.

Methods: A total of 722 adult patients with laboratory-confirmed COVID-19 pneumonia were analyzed in this single-center retrospective study between March 15 and May 1, 2020.

Results: A total of 722 laboratory-confirmed patients with COVID-19 pneumonia were included in the study. There were 235 (32.5%) elderly patients and 487 (67.5%) non-elderly patients. The most common comorbidities were hypertension (251 [34.8%]), diabetes mellitus (198 [27.4%]), and ischemic heart disease (66 [9.1%]). The most common symptoms were cough (512 [70.9%]), followed by fever (226 [31.3%]), and shortness of breath (201 [27.8%]). Lymphocytopenia was present in 29.7% of the patients, leukopenia in 12.2%, and elevated CRP in 48.8%. By the end of May 20, 648 (89.7%) patients had been discharged and 60 (8.5%) patients had died. According to our study, while our overall mortality rate was 8.5%, this rate was 14.5% in elderly patients, and the difference was significant.

Conclusions: This case series provides characteristics and outcomes of sequentially adult patients hospitalized with laboratory-confirmed COVID-19 pneumonia in Turkey.

© 2020 Japanese Society of Chemotherapy and The Japanese Association for Infectious Diseases.

Published by Elsevier Ltd. All rights reserved.

1. Introduction

Coronavirus disease 2019 (COVID-19) was first identified in Wuhan, China, on December 31, 2019. COVID-19 is a respiratory illness caused by Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2). Unfortunately, SARS-CoV-2 had quickly spread to the other countries and resulted in a pandemic. The number of infected patients continues to rise dramatically. Globally, as June 17, 2020, there have been 8,061,550

confirmed cases of COVID-19, including 440,290 deaths, reported to WHO [1-4].

In Turkey, the first case of COVID-19 was reported by the Ministry of Health of the Republic of Turkey on March 10, 2020. Although this was somewhat later than in European countries, as of June 17, 2020, the total number of COVID-19 cases and deaths reached 181,298 and 4842 respectively on. Our country has managed to slow the spread of the virus as a result of the very strict measures we have followed during the last two months [4].

E-mail address: elifsarginaltunok@gmail.com (E.S. Altunok).

^{*} Corresponding author.

The clinical spectrum of COVID-19 has a great variation from asymptomatic infection to acute respiratory distress syndrome and eventually death. The mortality rates vary across the countries probably due to the heterogeneity in study characteristics and patient cohorts as well as treatment strategies. As there is not currently approved drug to treat COVID-19, there were great variations in the mortality of COVID-19 globally even in the same city [3,5–7]. Therefore, we aimed to summarize the clinical characteristics and outcomes of 722 adult patients hospitalized with laboratory-confirmed COVID-19 pneumonia in Istanbul, Turkey.

2. Methods

2.1. Study design and participants

This single-center retrospective study was approved by the Ethics Committee of Istanbul Gaziosmanpasa Training and Research Hospital, and the requirement for written informed consent was waived by our ethics committee. Our hospital has served as a pandemic hospital during the outbreak in Istanbul, Turkey.

A total of 1153 adult patients (≥18 years old) with suspected or laboratory-confirmed COVID-19 pneumonia were hospitalized between March 15 and May 1, 2020. All patients had a fever or other respiratory symptoms with manifestations of pneumonia. Of these patients, 722 (62.6%) had a positive nasopharyngeal real-time polymerase chain reaction (RT-PCR) result for COVID-19 with chest computed tomography (CT) findings compatible with COVID-19 pneumonia were included in this retrospective study. All patients were followed up to death or discharge or the end of the study (May 20, 2020).

2.2. Data collection

Demographic, clinical findings, laboratory results, radiological features, treatments (antiviral, antibacterial, systemic corticosteroid, tocilizumab, and respiratory support), and outcomes were obtained from electronic medical records of patients. We divided patients into two groups according to age, as an elderly group (\geq 65 years old) and non-elderly group (<65 years old). All raw data were initially evaluated by trained physicians. The outcomes were discharge, ICU admission, mortality, and risk factors for mortality.

2.3. CT technique and image interpretation

Among the 722 patients, CT scans of 65 patients were performed in another hospital. Thus we were able to evaluate CT scans of 657 patients. The chest CT scans were obtained using the standard dose protocol of our hospital with a 128-slice multi-detector CT scanner (Optima; General Electric Healthcare, Wisconsin, USA). All CT scans were performed during a single breath-hold without contrast administration. All CT images were reviewed by a radiologist with 9 years of experience in interpreting chest CT imaging (FC), on a PACS imaging workstation (Infinitt PACS; Infinitt Healthcare, Seoul, Korea).

The CT features that were evaluated included; ground-glass opacities (GGO), consolidation, crazy paving pattern, tree-in-bud sign, air bronchogram, subpleural linear opacity, halo and reversed halo signs. The terms were defined in accordance with the Fleischner Society guidelines [8]. The location of the lesion was classified as predominantly central or predominantly peripheral, depending on whether it was found in the inner or outer half of the lung field, respectively. The affected lung and lobes pleural and pericardial effusion, the presence of mediastinal lymphadenopathy (short axis>1 cm) and bronchiectasis were also noted.

2.4. Definitions

Disease severity on admission was defined on the basis of COVID-19 Diagnosis and Treatment Guide' published by The Ministry of Health of the Republic of Turkey [9]. The patients were categorized into three groups according to their disease severity. The severe illness was defined as the presence of one of the followings: (a) respiratory distress with respiratory frequency \geq 30/min; (b) pulse oximeter oxygen saturation at rest <93%; and (c) artery partial pressure of oxygen/inspired oxygen fraction, PaO2/FiO2) \leq 300 mm Hg. Critical illness was defined as the requirement of high flow oxygen or non-invasive or invasive mechanical ventilation. The other patients who did not meet the above criteria were classified as having the non-severe illness.

2.5. Treatment

In Turkey, our treatment options for COVID-19 include Hydroxychloroquine (200 mg every 12 h, orally, 5–10 days), Favipiravir (first day 1600 mg, and then 600 mg every 12 h, orally, for 5–7 gün) and Lopinavir-ritonavir (500 mg twice daily, orally, for 10–14 days). Most patients have received a combination of hidroksiqlorokin and azithromycin (500 mg every 24 h, orally, for 5 days). The severe and critically ill patients were received with Favipiravir or Lopinavir-ritonavir based on the COVID-19 Diagnosis and Treatment Guide' published by The Ministry of Health of the Republic of Turkey [10]. Oseltamivir (75 mg every 12 h, orally, for 5–10 days) was also added to the current treatment during the influenza season. In addition, methylprednisolone for 3–15 days and Tocilizumab were received in seriously ill patients in a cytokine storm.

2.6. Statistics analysis

The data obtained were analyzed using an IBM SPSS Statistics 25 program and checked for suitability for a normal distribution with the Shapiro-Wilk test. Categorical variables were presented as counts and percentages. Continuous variables were presented as mean and standard deviation (SD), otherwise as the median and interquartile range (IQR). Continuous variables if normally distributed were analyzed by independent sample *t*-test; otherwise, the Mann-Whitney *U* test was used. All categorical data were analyzed with the chi-square test. Variables significantly associated with mortality in univariate analysis, analyzed in multivariate logistic regression analysis to determine independent risk factors for mortality. P values <0.05 indicated that the difference was statistically significant.

3. Results

3.1. Characteristics of patients at admission

3.1.1. All patients

A total of 722 laboratory-confirmed patients with COVID-19 pneumonia were included in the study. The mean age was 57.5 ± 15.3 years (range, 18-108 years), and 371 of 722 (51.4%) were male. In total, 183 (25.3%) patients had only one comorbidity, and 215 (29.8%) patients had more than one comorbidities. The most common comorbidities were hypertension (251 [34.8%]), diabetes mellitus (198 [27.4%]), and ischemic heart disease (66 [9.1%]) (Table 1). The most common symptoms were cough (512 [70.9%]), followed by fever (226 [31.3%]), and shortness of breath (201 [27.8%]). Less common symptoms included muscle ache, gastrointestinal system (diarrhea, nausea, and vomiting) symptoms, and headache. At triage, 129 patients (17.9%) were febrile (>38 °C), 29

Table 1Baseline characteristics of adult patients hospitalized with laboratory-confirmed COVID-19 pneumonia.

Characteristic	All patients ($n = 722$)	Non-elderly patients ($n=487$)	Elderly patients ($n=235$)	P-value ^a	
Age, (mean ± SD), years	57.5 ± 15.3	49.3 ± 10.5	74.5 ± 7.7	<0.001	
Male, n (%)	371 (51.4)	262 (53.8)	109 (46.4)	0.068	
Comorbidities, n (%)					
None	324 (44.9)	281 (57.7)	43 (18.3)	< 0.001	
1	183 (25.3)	111 (22.8)	72 (30.6)	0.028	
>1	215 (29.8)	95 (19.5)	120 (51.1)	< 0.001	
Hypertension	251 (34.8)	108 (22.2)	143 (60.9)	< 0.001	
Diabetes mellitus	198 (27.4)	104 (21.4)	94 (40.0)	< 0.001	
Asthma	46 (6.4)	24 (4.9)	22 (9.4)	0.033	
Chronic obstructive pulmonary disease	28 (3.9)	11 (2.3)	17 (7.2)	0.002	
Hyperlipidemia	38 (5.3)	22 (4.5)	16 (6.8)	0.215	
Chronic kidney disease	27 (3.7)	15 (3.1)	12 (5.1)	0.209	
Congestive Heart Failure	21 (2.9)	6 (1.2)	15 (6.4)	< 0.001	
İschemic Heart Disease	66 (9.1)	29 (6.0)	37 (15.7)	< 0.001	
Neurologic disease	18 (2.5)	6(1.2)	12 (5.1)	0.004	
Rheumatological Disease	7 (1.0)	6 (1.2)	1 (0.4)	0.437	
Malignancy	12 (1.7)	6 (1.2)	6 (2.6)	0.219	
Other comorbidities ^b	26 (3)	10 (2)	16 (5)	0.001	
Symptoms, n (%)	(_)	(-)	(-)		
More than one symptom	346 (47.9)	235 (48.3)	111 (47.2)	0.812	
Fever	226 (31.3)	159 (32.6)	67 (28.5)	0.267	
Cough	512 (70.9)	355 (72.9)	157 (66.8)	0.097	
Shortness of breath	201 (27.8)	127 (26.1)	74 (31.5)	0.133	
Chest pain	5 (0.7)	4 (0.8)	1 (0.4)	1.000	
Muscle ache	77 (10.7)	55 (11.3)	22 (9.4)	0.520	
Headache	28 (3.9)	17 (3.5)	11 (4.7)	0.420	
Sore throat	18 (2.5)	13 (2.7)	5 (2.1)	0.802	
Gastrointestinal System Symptoms ^c	47 (6.5)	33 (6.8)	14 (6.0)	0.749	
Onset of symptoms to hospitalization (days)	4 (2-7)	5 (2-7)	3 (0-7)	0.082	
Initial vitals, n (%)	1(2 /)	3 (2 7)	3 (0 7)	0.002	
Temperature >38 °C	129 (17.9)	99 (20.3)	30 (12.8)	0.013	
Oxygen saturation <90%	150 (20.8)	79 (16.2)	71 (30.2)	< 0.001	
Respiratory rate >30 breaths/min	29 (4.0)	18 (3.7)	11 (4.7)	0.547	
Heart rate >100 beats/min	19 (2.6)	10 (2.1)	9 (3.8)	0.213	
Systolic arterial pressure <90 mmHg, or diastolic <60 mmHg	12 (1.7)	3 (0.6)	9 (3.8)	0.003	
Oxygen support at admission, n (%)	12 (1.7)	3 (0.0)	3 (3.0)	0.003	
Nasal cannula	108 (15)	57 (11.7)	51 (21.7)	< 0.001	
Face mask	32 (4.4)	18 (3.7)	14 (6)	0.16	
Non-invasive mechanical ventilation	2 (0.3)	1 (0.2)	1 (0.4)	0.10	
Invasive mechanical ventilation	8 (1.1)	3 (0.6)	5 (2.1)	0.06	
Clinical severity assessment, n (%)	0 (1.1)	3 (0.0)	3 (2.1)	0.00	
Non-severe illness	543 (75.2)	390 (80.1)	153 (65.1)	< 0.001	
Severe illness	90 (12.5)	51 (10.5)	39 (16.6)	<0.001 0.02	
Critically illness	89 (12.3)	43 (8.8)	` '	<0.02 <0.001	
Citically lilless	09 (12.3)	43 (0.6)	46 (19.6)	<0.001	

Abbreviations: SD, standard deviation; NA, not applicable.

Those with significant p values are indicated in bold (P values < 0.05 indicated that the difference was statistically significant).

(4.0%) had a respiratory rate greater than 30 breaths/minute, and 150 (20.8%) had oxygen saturation less than 90% (Table 1). The median leucocytes count was 6.2 ($\times 10^3 / \mu L$), the neutrophil count was 4.3 ($\times 10^3 / \mu L$), lymphocyte count was 1.3 ($\times 10^3 / \mu L$), C-reactive protein (CRP) was 37.5 (mg/L), procalcitonin was 0.16 (ng/mL), and ferritin was 171 (ng/mL). Lymphocytopenia was present in 29.7% of the patients, leukopenia in 12.2%, and elevated CRP in 48.8% (see Table 2).

At admission, the CT findings of the patients are shown in Table 3. Of these 657 patients, 87% of patients had involvement of two or more lobes, 87% of lesions were located mainly in the peripheral zone of the lung. When a single lobe was involved, the right lower lobe was most often affected (13/36 [36%]). The most common CT features were patchy or rounded GGO (51%) and GGO with consolidation (39%). Seventy-three (11%) patients had the crazy-paving pattern, 194 (30%) had subpleural linear opacity. The air bronchogram sign was visualized in 216 (33%) patients, the halo sign in 131

(20%), and the reversed halo sign in 60 (9%). The CT findings and other data are presented in Table 3.

Clinical severity assessment of COVID-19 pneumonia was defined in 3 groups. The distribution of clinical severity was 543 (75.2%), 90 (12.5%), and 89 (12.3%) for non-severe, severe, and critical respectively (Table 1).

3.1.2. Elderly vs non-elderly patients

There were 235 (32.5%) elderly patients and 487 (67.5%) non-elderly patients. The mean age of elderly patients was 74.5 years (SD, \pm 7.7; range, 65–108 years), and 109 of 235 (46.4%) elderly patients were male. In elderly patients, 72 (30.6%) patients had only one comorbidity, and 120 (51.1%) patients had more than one comorbidities. The prevalence of more than one comorbidity was significantly higher among elderly patients. The most common comorbidities were hypertension (60.9%), diabetes mellitus (40%), and ischemic heart disease (15.7%). Elderly patients compared to

^a P values indicate differences between elderly and non-elderly patients.

^b Other comorbidities: Atrial fibrillation, hypothyroidism, benign prostatic hyperplasia, solid organ transplantation, chronic hepatitis B infection, chronic hepatitis c infection and liver cirrhosis.

^c Gastrointestinal System Symptoms: Diarrhea, nausea and vomiting.

Table 2 Initial laboratory test results of adult patients hospitalized with laboratory-confirmed COVID-19 pneumonia.

Parameter Median (IQR)	All patients, $(n=722)$	Non-elderly patients (n $=$ 487)	Elderly patients (n $= 235$)	P-value ^a
Leucocytes, × 103/μL (normal range 4.1–11)	6.2 (4.9-8.1)	5.9 (4.8-7.7)	6.6 (5.2–8.5)	0.001
Leucocytes <4.1	88 (12.2)	67 (13.8)	21 (8.9)	0.06
Neutrophils, $\times 10^3/\mu L$ (normal range 2–8)	4.3 (3.1-5.9)	4.1 (2.9-5.6)	4.5 (3.4-6.5)	< 0.001
Lymphocytes, $\times 10^3/\mu$ L (normal range 1–5)	1.3 (9.5-1.7)	1.3 (9.6-1.8)	1.2 (8.8-1.7)	0.053
Lymphocytes <1	211 (29.7)	136 (28.3)	75 (32.5)	0.29
Haemoglobin, g/dL (normal range 11.5-15)	13.4 (12.5-14.4)	13.6 (12.7-14.5)	13 (12-14.2)	< 0.001
Platelets, × 103/μL (normal range 150–400)	195 (159-241)	193 (158-239)	199 (162-245)	0.47
Aspartate aminotransferase, U/L (normal range 0-35)	29 (23-41)	29 (23-41)	30 (22-42)	0.76
Alanine aminotransferase, U/L (normal range 0-35)	23 (17-35)	25 (18-38)	20 (14-29)	< 0.001
Urea, mg/dL (normal range 17-43)	32 (24-44)	28 (22-37)	41 (32-58.5)	< 0.001
Serum creatinine, mg/dL (normal range 0.51-0.95)	0.84 (0.67-1.05)	0.82 (0.65-0.98)	0.92 (0.75-1.33)	< 0.001
Activated partial thromboplastin time, sn (normal range 16.9–31.9)	25.4 (24-27.1)	25.3 (24-27)	25.5 (23.9-27.4)	0.49
Prothrombin time, sec (normal range 9.4–14.2)	12.8 (12.1-13.8)	12.8 (12.1-13.7)	13 (12.1-14.2)	0.03
C-reactive protein, mg/L (normal range 0-5)	37.5 (13-90.7)	33 (12-85)	48 (18-103)	0.02
C-reactive protein >40	330 (48.8)	212 (45.8)	118 (55.4)	0.02
Lactate dehydrogenase, U/L (normal range 0-247)	271 (217-351)	277 (216-350)	261 (217-353)	0.77
Ferritin, ng/mL (male: 20-250 female: 10-120)	171 (85-375)	174 (80-378)	170 (90-393)	0.54
Procalcitonin, ng/mL	0.16 (0.12-0.24)	0.15 (0.12-0.22)	0.18 (0.12-0.34)	0.009
Procalcitonin >0.5	49 (13.4)	25 (10.6)	24 (18.5)	0.03
High sensitivity troponin I, ng/L (<11.6)	4.1 (2.4-9.1)	3.4 (2-5.5)	7.8 (4.4-17.7)	< 0.001
CK-MB, ng/mL (normal range 0.6-6.3)	1 (0.7-1.9)	0.9 (0.6-1.5)	1.5 (0.9-2.8)	< 0.001
Fibrinogen, mg/dL (normal range 200-400)	355 (313-401)	353 (316-405)	361 (309-401)	0.97

Abbreviations: NA, not applicable; IQR, interquartile range.

Those with significant p values are indicated in bold (P values < 0.05 indicated that the difference was statistically significant).

non-elderly patients had higher rates of hypertension, diabetes mellitus, and ischemic heart disease, the difference was statistically significant. The most common symptoms were cough (66.8%), followed by shortness of breath (31.5%), and fever (28.5%). At triage, 30 elderly patients (12.8%) were febrile (>38 $^{\circ}$ C), 11 (4.7%) had a respiratory rate greater than 30 breaths/minute, and 71 (30.2%) had oxygen saturation less than 90%. At admission, elderly patients had less fever and low oxygen saturation was more common. The median leucocytes count was 6.6 ($x10^3/\mu L$), the neutrophil count was 4.5 (x10 3 / μ L), lymphocyte count was 1.2 (x10 3 / μ L), CRP was 48 (mg/ L), procalcitonin was 0.18 (ng/mL), and ferritin was 170 (ng/mL). Lymphocytopenia was present in 32.5% of the patients, leukopenia in 8.9%, and elevated CRP in 55.4%. The distribution of clinical severity was 39 (16.6%), and 46 (19.6%) for severe and, critical respectively which was significantly higher than non-elderly patients (Table 1).

3.2. Treatments and outcomes

3.2.1. All patients

At admission, 712 of 722 patients were admitted to the ward and 10 patients were admitted to the intensive care unit. At ward, oxygen therapy was applied in 108 (15%) patients with the nasal cannula and 32 (4.4%) patients with face mask at admission (Table 1). 79 of 712 patients during the follow up at ward were transferred to ICU. The median duration from the onset of hospitalization to ICU admission was 3 days (IQR, 1.7–6) (Table 4). Overall, 89 (12.3) patients were admitted to the intensive care unit. In ICU, the number of patients required high flow oxygen support, non-invasive mechanical ventilation, and invasive mechanical ventilation were 19 (2.6%), 10 (1.4%), and 60 (8.3%) respectively (Table 4).

By the end of May 20, 648 (89.7%) patients had been discharged and 60 (8.5%) patients had died; all other patients were still hospitalized. The median hospital duration was 6 [4—10] days. 12 of 14 patients still being hospitalized are followed up at the ward and 2 at the ICU by the end of May 20 (Table 4). Finally, on the multivariable analysis, older age and elevated CRP remained the significant independent risk factors for death (Table 5).

3.2.2. Elderly vs non-elderly patients

At admission, 229 of 235 elderly patients were admitted to the ward and 6 elderly patients were admitted to the intensive care unit. At the ward, oxygen therapy was applied in 51 (21.7%) patients with the nasal cannula and 14 (6%) patients with face mask at admission (Table 1). 40 of 229 elderly patients during the follow up at ward were transferred to ICU. The median duration from the

Table 3CT Features of adult patients hospitalized with laboratory-confirmed COVID-19 pneumonia.

meuniona.				
Variables	All patients (n = 657)			
CT findings, n (%)				
GGO	335 (51%)			
GGO with consolidation	255 (39%)			
Consolidation	6 (1%)			
Crazy paving pattern	73 (11%)			
Tree-in-bud sign	13 (2%)			
Air bronchogram	216 (33%)			
Subpleural linear opacity	194 (30%)			
Halo sign	131 (20%)			
Reversed halo sign	60 (9%)			
Pleural effusion	32 (5%)			
Pericardial effusion	8 (1%)			
Mediastinal lymphadenopathy	60 (9%)			
Lung Involvement, n (%)				
Right lung	46 (7%)			
Left lung	14 (%2)			
Bilateral	595 (91%)			
Distribution of lesions in lung, n (%)				
Peripheral dominance	569 (87%)			
Central dominance	9 (1%)			
Peripheral and central	77 (12%)			
Upper lobes dominance	83 (13%)			
Lower lobes dominance	403 (61%)			
Upper and lower lobe	169 (26%)			
Frequency of lobe involvement, n (%)				
Right upper lobe	545 (83%)			
Right middle lobe	500 (76%)			
Right lower lobe	612 (93%)			
Left upper lobe	552 (84%)			
Left upper lobe	586 (89%)			
More than two lobes affected	573 (87%)			

Abbreviation: GGO, ground-glass opacities.

^a P values indicate differences between elderly and non-elderly patients.

Table 4 Clinical outcomes of adult patients hospitalized with laboratory-confirmed COVID-19 pneumonia.

Characteristic of hospitalization	All patients, $(n = 722)$	Non-elderly patients ($n = 487$)	Elderly patients ($n = 235$)	P-value ^a
Median hospital duration, days	6 (4–10)	5 (3-9)	8 (5–12)	<0.001
Ventilatory support at ICU, n (%)				
High flow oxygen	19 (2.6)	6 (1.2)	13 (5.5)	0.001
Non-invasive mechanical ventilation	10 (1.4)	7 (1.4)	3 (1.3)	0.86
Invasive mechanical ventilation	60 (8.3)	30 (6.2)	30 (12.8)	0.003
Onset of hospitalization to ICU admission, days	3 (1.7–6)	3 (1-6)	3 (2-6)	0.24
Admitted to ICU, n (%)	89 (12.3)	43 (8.8)	46 (19.6)	< 0.001
Discharged, n (%)	648 (89.7)	454 (93.2)	194 (82.6)	< 0.001
Died, n (%)	60 (8.5)	26 (5.3)	34 (14.5)	< 0.001
Still hospitalized, n (%)	14 (1.9)	7 (1.4)	7 (3.0)	0.16
in ward	12 (1.7)	6 (1.2)	6 (2.6)	0.19
in ICU	2 (0.3)	1 (0.2)	1 (0.4)	0.6

Abbreviation: ICU, intensive care unit.

Those with significant p values are indicated in bold (P values < 0.05 indicated that the difference was statistically significant).

Table 5Independent risk factors for mortality in patients with laboratory-confirmed COVID-19 pneumonia.

Parameter	OR	CI (95%)		p value
		Lower	Upper	
Age (years)	1.04	1.007	1.082	0.02
C-reactive protein, mg/L	1.01	1.012	1.024	< 0.001
Ferritin (ng/mL)	1.00	1.000	1.001	0.10
Diabetes mellitus (+)	1.97	0.718	5.415	0.18
Hypertension (+)	0.75	0.263	2.153	0.59

Those with significant p values are indicated in bold (P values < 0.05 indicated that the difference was statistically significant).

onset of hospitalization to ICU admission was 3 days (IQR, 2–6) (Table 4). Overall, 46 (19.6) patients were admitted to the intensive care unit. In ICU, the number of elderly patients required high flow oxygen support, non-invasive mechanical ventilation, and invasive mechanical ventilation were 13 (5.5%), 3 (1.3%), and 30 (12.8%) respectively (Table 4).

By the end of May 20, 194 (82.6%) patients had been discharged and 34 (14.5%) patients had died; all other patients were still hospitalized. The median hospital duration was 8 [5–12] days. 6 of 7 patients still being hospitalized are followed up at the ward and 1 patients at the ICU by the end of May 20 (Table 4).

4. Discussion

A total of 722 adult patients with laboratory-confirmed COVID-19 pneumonia was analyzed in this single-center retrospective study between March 15 and May 1, 2020. We identified major clinical characteristics, laboratory results, radiological features, and outcomes for the disease. We divided patients into two groups according to age, as the elderly group and non-elderly group. We also identified independent risk factors for mortality. Previous reports described different mortality rates for death in adults who were hospitalized with COVID-19 among countries. Mortality rates in studies reported from China vary between 1.4 and 12.8 [10-12]. In the first large case series from the US, mortality rates were reported as 21% [13]. In a study conducted in Italy, the mortality rate in inpatients was reported as 20.6% [14]. According to our study, while our overall mortality rate was 8.5%, this rate was 14.5% in elderly patients, and the difference was significant. Compared to other countries, these mortality rates were significantly lower than the US and Italy and was similar to data reported from China.

Globally, as June 17, 2020, there have been 8,061,550 confirmed cases of COVID-19, including 440,290 deaths, reported to WHO [4]. Although most patients present with mild symptoms that are not

life-threatening, the number of deaths is still high owing to the large patients' population. In the previous studies identified several risk factors for death in adults who were hospitalized with COVID-19. In particular, older age, d-dimer levels greater than 1 μ g/mL, elevated levels of blood IL-6, high-sensitivity cardiac troponin I, and lactate dehydrogenase and lymphopenia were more commonly seen in severe COVID-19 illness in-hospital death [15]. Our study confirmed that increased age was associated with death in patients with COVID-19. In addition, those with hypertension and diabetes mellitus were highly prevalent in this case series, but this was not related to the higher rate of comorbidity in elderly patients.

COVID-19 is a viral disease characterized by decreased lymphocyte count. According to our current information, cytokine storm plays an important role in severe COVID-19 cases. SARS-CoV-2 is known to mainly affect lymphocytes, especially T lymphocytes, and virus particles induce a cytokine storm in the body, this results in lymphopenia [3,16]. In our study, the absolute value of lymphocytes in most patients decreased (29.7%). However, lymphopenia was not associated with mortality in the analyzes. Among laboratory abnormalities, only elevated CRP remained the significant independent risk factors for death in our study.

Because of the primary involvement of the respiratory system, chest CT is strongly recommended in suspected COVID-19 cases, for initial evaluation. The characteristic patterns and distribution of initial CT manifestations in COVID-19 cases include bilateral, multilobar ground-glass opacification with a peripheral or posterior distribution (or both), mainly in the lower lobes [17,18]. According to our study, chest CT showed similar characteristics in the majority of patients, such as ground-glass opacification 335 (51%), bilateral involvement 595 (91%), peripheral distribution 569 (87%), and multilobar (more than two lobes) involvement 573 (87%). Pleural effusion, pericardial effusion, cavitation, pneumothorax, and lymphadenopathy are some of the uncommon but possible findings seen with disease progression [19,20]. In our study, halo sign (20%) and air bronchogram (33%) findings were found to be relatively high on admission.

Istanbul is an overpopulated city that can be seen as a reflection of Turkey by having significant citizen diversity as a result of being a migration receiving city. At the same time, this city has been the center of COVID-19 pandemic. During the Covid-19 pandemic, the majority of cases in our country were placed in this city. Therefore it can be said that it is important in terms of the data reflect the turkey.

5. Limitations

This study has several limitations. First of all, it shows the results of a single-center in Istanbul, Turkey. Second, Due to the

^a P values indicate differences between elderly and non-elderly patients.

retrospective nature of the study, the missing data were collected from the patients' electronic medical records. This precluded the level of detail possible with a manual medical record review. Thirdly, some laboratory tests (for example, D-Dimer, IL-6) were not done in all the patients, and missing data or important tests might lead to bias of clinical characteristics.

6. Conclusions

This case series provides characteristics and outcomes of sequentially adult patients hospitalized with laboratory-confirmed COVID-19 pneumonia in Istanbul, Turkey. In addition, it reveals risk factors associated with mortality.

Contributors

Elif SARGIN ALTUNOK was responsible for the organization and coordination of the trial. Elif SARGIN ALTUNOK was the chief investigator, and responsible for the data analysis with Celal Satici and Mustafa Asim Demirkol. Elif SARGIN ALTUNOK, Celal Satici, Mustafa Asim Demirkol, Mustafa Alkan, Bengul Gursoy and Ferhat Cengel developed the trial design. All authors contributed to the writing of the final manuscript. All authors contributed to the management or administration of the trial.

Funding

There is no source of funding.

Declaration of competing interest

The authors declared that they have no conflicts of interest.

Acknowledgments

Our sincere thanks to all healthcare professionals for their brave efforts in COVID-19 treatment, prevention, and control.

References

- [1] Lu H, Stratton CW, Tang YW. Outbreak of pneumonia of unknown etiology in Wuhan China: the mystery and the miracle. J Med Virol 2020. https://doi.org/10.1002/jmv.25678. published online Jan 16.
- [2] Zhu N, Zhang D, Wang W, Li X, Yang B, Song J, et al. A novel coronavirus from patients with pneumonia in China, 2019. N Engl J Med 2020;382(8):727–33. https://doi.org/10.1056/NEJMoa2001017.
- [3] 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):507–13. https://doi.org/10.1016/S0140-6736(20)30211-7. Feb 15.
- [4] World Health Organization. Coronavirus disease (COVID-19) dashboard. Available from: https://covid19.who.int/; 2020.
- [5] Li X, Xu S, Yu M, Wang K, Tao Y, Zhou Y, et al. Risk factors for severity and mortality in adult COVID-19 inpatients in Wuhan. J Allergy Clin Immunol 2020 Apr 12. https://doi.org/10.1016/j.jaci.2020.04.006. S0091-6749(20)30495-4.
- [6] Jan H, Faisal S, Khan A, Khan S, Usman H, Liaqat R, et al. COVID-19: review of epidemiology and potential treatments against 2019 novel coronavirus. Discoveries (Craiova) 2020 Apr 26;8(2):e108. https://doi.org/10.15190/d.2020.5.
- [7] Zhang G, Hu C, Luo L, Fang F, Chen Y, Li J, et al. Clinical features and short-term outcomes of 221 patients with COVID-19 in Wuhan, China. J Clin Virol 2020 Jun;127:104364. https://doi.org/10.1016/j.jcv.2020.104364.
- [8] Hansell DM, Bankier AA, MacMahon H, McLoud TC, Müller NL, Remy J. Fleischner Society: glossary of terms for thoracic imaging. Radiology 2008:246:697–722.
- [9] The Ministry of Health of the Republic of Turkey. COVID-19 diagnosis and treatment Guide. https://covid19.saglik.gov.tr/TR-66337/genel-bilgilerepidemiyoloji-ve-tani.html.
- [10] Liu K, Fang YY, Deng Y, Liu W. Clinical characteristics of novel coronavirus cases in tertiary hospitals in Hubei province. Chin Med J (Engl) 2020 May 5;133(9):1025–31. https://doi.org/10.1097/CM9.00000000000000744.
- [11] Chen TL, Dai Z, Mo P, Li X, Ma Z, Song S, et al. Clinical characteristics and outcomes of older patients with coronavirus disease 2019 (COVID-19) in Wuhan, China (2019): a single-centered, retrospective study. J Gerontol A Biol Sci Med Sci 2020 Apr 11. https://doi.org/10.1093/gerona/glaa089. glaa089.
- [12] Guan WJ, Ni ZY, Hu Y, Liang WH, Ou CQ, He JX, et al. Clinical characteristics of coronavirus disease 2019 in China. N Engl J Med 2020 Apr 30;382(18): 1708–20. https://doi.org/10.1056/NEJMoa2002032.
- [13] Richardson S, Hirsch JS, Narasimhan M, Crawford JM, McGinn T, Davidson KW, et al. Presenting characteristics, comorbidities, and outcomes among 5700 patients hospitalized with COVID-19 in the New York city area. JAMA 2020;323(20):2052–9. https://doi.org/10.1001/jama.2020.6775. Apr 22.
- [14] Giacomelli A, Ridolfo AL, Milazzo L, Oreni L, Bernacchia D, Siano M, et al. 30-day mortality in patients hospitalized with COVID-19 during the first wave of the Italian epidemic: a prospective cohort study. Pharmacol Res 2020 May 22;158:104931. https://doi.org/10.1016/j.phrs.2020.104931.
 [15] Zhou F, Yu T, Du R, Fan G, Liu Y, Liu Z, et al. Clinical course and risk factors for
- [15] Zhou F, Yu T, Du R, Fan G, Liu Y, Liu Z, et al. Clinical course and risk factors for mortality of adult inpatients with COVID-19 in Wuhan, China: a retrospective cohort study. Lancet 2020 Mar 28;395(10229):1054–62. https://doi.org/ 10.1016/S0140-6736(20)30566-3.
- [16] Liu WJ, Zhao M, Liu K, Xu K, Wong G, Tan W, et al. T-cell immunity of SARS-CoV: implications for vaccine development against MERS-CoV. Antivir Res 2017:137:82–92.
- [17] Jin YH, Cai L, Cheng ZS, Cheng H, Deng T, Fan YP, et al. Zhongnan hospital of Wuhan university novel coronavirus management and Research team; evidence-based medicine chapter of China international exchange and promotive association for medical and Health care (CPAM). A rapid advice guideline for the diagnosis and treatment of 2019 novel coronavirus (2019-nCoV) infected pneumonia (standard version). Mil Med Res 2020;7:4.
- [18] Salehi S, Abedi A, Balakrishnan S, Gholamrezanezhad A. Coronavirus disease 2019 (COVID-19): a systematic review of imaging findings in 919 patients. AJR Am J Roentgenol 2020 Mar 14:1–7. https://doi.org/10.2214/AJR.20.23034.
- [19] Xu X, Yu C, Qu J, Zhang L, Jiang S, Huang D, et al. Imaging and clinical features of patients with 2019 novel coronavirus SARS-CoV-2. Eur J Nucl Med Mol Imag 2020 May;47(5):1275–80. https://doi.org/10.1007/s00259-020-04735-
- [20] Bernheim A, Mei X, Huang M, Yang Y, Fayad ZA, Zhang N, et al. Chest CT findings in coronavirus disease-19 (COVID-19): relationship to duration of infection. Radiology 2020 Jun;295(3):200463. https://doi.org/10.1148/ radiol.2020200463.