COVID-2019 Pneumonia Severity and distribution of lung changes observed on the initial chest X-ray as an indicator of final outcomes

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ABSTRACT: *Objectives:* This study aimed to assess the correlation between the severity of the initial chest x-ray (CXR) abnormalities in patients with a confirmed diagnosis of COVID-19 and the final outcomes. *Methods:* This retrospective study was conducted at the Royal Hospital, Oman between mid-March and May 2020 and included patients who had been admitted with a confirmed diagnosis of COVID-19 and had a final outcome. Serial CXRs were identified and examined for presence, extent, distribution and progression pattern of radiological abnormalities. Each lung field was divided into three zones on each CXR and a score was allocated for each zone (0 is normal and 1–4 is mild–severe). The scores for all six zones per CXR examination were summed to provide a cumulative chest radiographic score (range: 0–24). *Results:* A total of 64 patients were included; the majority were male (89.1%) and the mean age was 50.22 ± 14.86 years. The initial CXR was abnormal in 60 patients (93.8%). The most common finding was ground glass opacity (n = 58, 96.7%) followed by consolidation (n = 50, 83.3%). Most patients had bilateral (n = 51, 85.0%), multifocal (n = 57, 95.0%) and mixed central and peripheral (n = 36, 60.0%) lung abnormalities. The median score of initial CXR for deceased patients was significantly higher than recovered patients (17 versus 11; *P* = 0.009). Five CXR evolution patterns were identified: type I (initial radiograph deteriorates then improves), type II (fluctuate), type III (static), type IV (progressive deterioration) and type V (progressive improvement). *Conclusion:* A higher baseline CXR score is associated with higher mortality rate and poor prognosis in those with COVID-19 pneumonia.

Keywords: SARS-CoV-2; COVID-19; X-ray Film; Pneumonia; Prognosis; Oman.

Advances in Knowledge

- The findings of this study demonstrate that severe lung abnormalities on chest radiographs are associated with poor clinical outcome prognosis and increased mortality.

Application to Patient Care

- This study highlights the importance of chest radiographs in the initial diagnosis, follow-up and prediction of clinical course/outcome of patients with COVID-19.

N WUHAN, CHINA IN DECEMBER 2019, SEVERAL cases of severe pneumonia of unknown aetiology were admitted to the hospitals. By January 2020, the cause was identified as a novel coronavirus that was called severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) by the International Committee on Taxonomy of Viruses on February 22, 2020. The disease was named COVID-19 by the World Health Organisation (WHO) on the same day.^{1–3} Since then, the virus has spread worldwide and the WHO classified it as a pandemic on March 11, 2020.⁴

COVID-19 is diagnosed based on the clinical findings and real-time polymerase chain reaction (RT-PCR) assay of specimens obtained from the respiratory tract.^{5,6} Medical imaging, including chest computed tomography (CT) and chest x-rays (CXR), plays an important role in its diagnosis and disease progression monitoring upon admission.^{7–22}

In the Muscat Governorate of Oman, the Royal Hospital is the main referral hospital for the majority of patients with COVID-19 pneumonia who require admission; CXR is the main imaging modality for diagnosis and follow-up. Quantification of radiographic findings at initial presentation and follow-up can help determine the appropriate management of patients with COVID-19 pneumonia. Therefore, this study aimed to quantify the severity of the CXR findings using a scoring system and to examine the relationship between the severity of the lung changes on the baseline CXR and the final outcomes to determine its prognostic value. Secondly, this study aimed to evaluate the evolution pattern of follow-up CXRs of patients admitted with COVID-19 pneumonia.

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Methods

This retrospective study included all patients with confirmed diagnosis of COVID-19 who were admitted to the Royal Hospital in Muscat, Oman, between mid-March and May 2020 and had a final outcome.

The COVID-19 diagnosis was made based on a positive RT-PCR for SARS-CoV-2 of specimens obtained from the nasopharynx and oropharynx of all patients. Medical records of all patients were reviewed and demographic characteristics, presenting symptoms, vital signs, comorbidities, serum reactive protein, white blood count (WBC) and estimated glomerular filtration rate were obtained. The duration of hospitalisation, admission to the intensive care unit (ICU), history of intubation and the final outcome (death or recovery) were also documented.

All patients had initial frontal CXRs on the day of admission and follow-up CXRs during hospitalisation, obtained in the posteroanterior (PA) projection for those who were able to stand and anteroposterior projection for patients not able to stand. All CXRs were acquired using machines with digital radiography systems using standard techniques (Mobilet Mira Max, Siemens Medical Solution Diagnostic Ltd. Europe, Dublin, Ireland; DRX-Revolution, Carestream Health, Rochester, New York, USA).

Baseline and follow-up CXRs were reviewed retrospectively in consensus by three radiologists with 8–15 years of experience, with all three unaware of final patient outcomes. All CXRs were viewed using a dedicated radiology picture archiving and communication system and were assessed for the presence, distribution and extent of lung abnormalities, including consolidation, ground glass opacity (GGO), cavitation and pleural effusion. Distribution was considered peripheral if the changes were predominantly in the lateral half of the lung fields and central if they were predominantly in the medial half.

For the assessment of the severity of lung changes, each lung field was divided into three equal zones. This was due to technical reasons as most of the CXRs were taken bedside for critically ill patients and it was difficult to identify some anatomical landmarks. Each zone was assigned a score from 0-4 based on the percentage of lung involved (0 = no abnormality, 1 = <25% of the zone involved, 2 = 25-50% involved, 3 = 51-75% involved and 4 = >75% involved). The scores for all six zones of each CXR examination were summed to provide a cumulative chest radiographic score (range, 0-24) [Figure 1]. Evolution patterns of lung changes on serial CXRs were classified into five patterns: I) progression of CXR findings followed

by an improvement; II) fluctuation of CXR findings with at least two CXR peaks and an intervening, mild improvement of more than 25% from the overall mean of CXR scores; III) static CXR findings with no peaks or improvement of more than 25% from the overall mean of CXR scores; IV) progressive deterioration of CXR scores; and V) progressive improvement of CXR scores.

Patients were divided into two groups according to the final outcome: the deceased group and the recovered group. Continuous and discrete data were summarised using the median and interquartile range (IQR). Categorical data were summarised using frequency and percentage. The Chi-squared test (Fisher's exact test/Likelihood Ratio) was used to test the association between the categorical variables. The Mann-Whitney U test was used to compare nonnormally distributed continuous and discrete variables. Spearman's rank correlation coefficient was used to assess the correlation between the non-normally distributed continuous variables. Kolmogorov-Smirnov tests were performed to check normal distribution of continuous variables between the groups. Demographic characteristics, CXR scores, comorbidities and other important clinical features were also compared between the groups. A multivariate binary logistic regression analysis was performed to determine the independent predictors of mortality. A P value <0.05 was considered statistically significant. All statistical analyses were performed using Statistical Package for the Social Sciences (SPSS), Version 25 (IBM Corp., Chicago, Illinois, USA).

The Scientific Research Committee at the Royal Hospital approved this retrospective, single institution study and waived informed consent (SRC# 59/2020).

Results

A total of 64 patients were included, 57 men and seven women. The mean age was 50.22 ± 14.86 years. Of those, 29 patients (45.3%) had no comorbidities, 16 patients (25.0%) had a single comorbidity and 19 patients (29.7%) had multiple comorbidities. The duration of hospitalisation ranged between 7–19.5 days with a median of 11.5 days. Of the 64 patients, 44 (68.8%) were admitted to the ICU, 31 (48.4%) required intubation, 17 (26.6%) patients died and 47 (73.4%) recovered. The median duration of ICU admission was 11 days (IQR: 5.3–14.8 days). The median duration of intubation was 10 days (IQR: 7–16.8 days). The fatality rate among patients admitted to the ICU was 38.6% (n = 17) [Table 1].

The initial CXRs were acquired at a median of seven days (IQR: 5–8 days) from the onset of

Table 1: Characteristics of patients with a confirmed diagnosis of COVID-19 who were admitted to the Royal Hospital,Oman between mid-March and May 2020 (N = 64)

Characteristic	n (%)
Gender	
Male	57 (89.1)
Female	7 (10.9)
Age in years	
Mean ± SD	50.22 ± 14.86
Median (min-max)	46.5 (22–78)
Symptom	
Fever	47 (73.4)
Runny nose	7 (10.9)
Cough	40 (62.5)
SOB	41 (64.1)
Headache	4 (6.3)
Vomiting	1 (1.6)
Nausea	4 (6.3)
Dizziness	1 (1.6)
Muscle ache/body ache	6 (9.4)
Malaise	0 (0)
Lethargy	8 (12.5)
Phalangeal discomfort/pain	11 (17.2)
Chest pain	5 (7.8)
Pleuritic chest pain	1 (1.6)
Abdominal pain	0 (0)
Diarrhoea	12 (18.8)
Haemoptysis	0 (0)
Anorexia	3 (4.7)
Comorbidity	
No comorbidity	29 (45.3)
Single comorbidity	16 (25)
Multiple comorbidity	19 (29.7)
Diabetes mellitus	27 (42.2)
Ischemic heart disease	2 (3.1)
Hyperlipidaemia	5 (7.8)
Chronic kidney disease	6 (9.4)
Hypertension	19 (29.7)
Duration of hospital stay and disease cour	rse
Median days of hospitalisation (IQR)*	11.5 (7.0–19.5)
Number of patients admitted to the ICU	44 (68.8)

Median days of ICU admission (IQR)	11 (5.3–14.8)
Number of patients intubated	31 (48.4)
Median days of intubation (IQR)	10 (7.0–16.8)
Final outcome	
Recovered (i.e. discharged)	47 (73.4)
Deceased	17 (26.6)

*Hospitalisation duration was from the day of the admission to the day of discharge or death.

SD = standard deviation; *SOB* = shortness of breath; *IQR* = interquartile range; *ICU* = intensive care unit.

Table 2: Radiological findings of patients with confirmeddiagnosis of COVID-19 included in this study (N = 64)

Finding	n (%)
Abnormal CXR at admission	60 (93.8)
Median days since symptoms started and the CXR was acquired (IQR)	7 (5–8)
Abnormal CXR findings (n = 60)	
Consolidation	50 (83.3)
GGO	58 (96.7)
Cavitation	0 (0)
Central	4 (6.7)
Peripheral	20 (33.3)
Mixed central and peripheral	36 (60.0)
Unilateral	9 (15.0)
Bilateral	51 (85.0)
Unifocal	3 (5.0)
Multifocal	57 (95.0)
Pleural effusion	2 (3.3)
Total score of the baseline CXR	
Mean ± SD	11.91 ± 7.84
Median (IQR)	12.5 (4-18)

CXR = chest *x*-ray; *SD* = standard deviation; *IQR* = interquartile range; *GGO* = ground glass opacity.

symptoms and were abnormal in 93.8% of the patients (n = 60). The most common finding was GGO (n = 58, 96.7%), followed by consolidation (n = 50, 83.3%). Pleural effusion was a rare finding (n = 2, 3.3%) and none of the patients had cavitation. Bilateral lung involvement (n = 51, 85.0%) was more common than unilateral involvement (n = 9, 15.0%). The right lower (n = 50, 83.3%) and left lower (n = 51, 85.0%) zones were more commonly affected than the right middle (n = 45, 75.0%), left middle (n = 42, 70.0%), right upper (n = 35, 58.3%) and left upper (n = 28, 46.7%) zones. Mixed central and peripheral distribution was the

Variable Total baseline CXR score					
	Median score (IQR)	P value*			
Age group in year	s				
20-29	10 (0.5–19.5)				
30–39	13 (4–16)				
40-49	14 (4–22)	0.206			
50-59	7.5 (2.5–11.5)	0.296			
60–69	13.5 (6.3–22.8)				
≥70	12 (5–17)				
Diabetes mellitus					
Yes	13 (7–19)	0.140			
No	11 (2.5–17.5)	0.149			
Hypertension					
Yes	11 (5–18)	0.707			
No	13 (4–18.5)	0.797			
Ischaemic heart disease					
Yes	6.5 (0–13)	0.389			
No	12.5 (4–18.3)	0.288			
Chronic kidney di	sease				
Yes	3.5 (0.8–7.3)	0.010 [†]			
No	13 (4.8–19)	0.010			
Hyperlipidaemia					
Yes	13 (3–24)	0.625			
No	12 (4–18)	0.625			
ICU admission					
Yes	14 (12–21)	0.0001			
No	2.5 (1-5)	0.0001			
Intubation					
Yes	16 (13–21)	0.0001			
No	5 (1.5–12.5)	0.0001			
Outcome					
Recovered	11 (4–16)	o coot			
Deceased	17 (12–22.5)	0.0091			

Table 3: Association between Initial Chest X-ray score

CXR = chest x-ray; IQR = interquartile range; ICU = intensive care unit. *Using Mann-Whitney U test. [†]Statistically significant.

predominant distribution (n = 36, 60.0%), followed by peripheral (n = 20, 33.3%) and central distribution (n = 4, 6.7%). Multifocal distribution was more common (n = 57, 95.0%) than unifocal distribution (n = 3, 5.0%) [Table 2].

Table 4: Association between patients	' clinical and laboratory
findings and final outcome	

Variable	Total	n (%)		P
		Recovered	Deceased	value
Gender				
Male	57	42 (73.7)	15 (26.3)	1 000
Female	7	5 (71.4)	2 (28.6)	1.000
Mean age in years ± SD		46.57 ± 13.57	60.29 ± 13.91	0.001*
Diabetes melli	itus			
Yes	27	16 (59.3)	11 (40.7)	0.04.4*
No	37	31 (83.8)	6 (16.2)	0.044*
Ischemic hear	t disease	2		
Yes	2	2 (100)	0 (0)	1 000
No	62	45 (72.6)	17 (27.4)	1.000
Hyperlipidaen	nia			
Yes	5	4 (80.0)	1 (20.0)	
No	59	43 (72.9)	16 (27.1)	1.000
Chronic kidne	ey diseas	e		
Yes	6	6 (100)	0 (0)	
No	58	41 (70.7)	17 (29.3)	0.182
Hypertension				
Yes	19	12 (63.2)	7 (36.8)	
No	45	35 (77.8)	10 (22.2)	0.236
WBC count				
Normal	44	36 (81.8)	8 (18.2)	
Abnormal	20	11 (55.0)	9 (45.0)	0.034*
CRP in mg/L				
<10	1	1 (100)	0 (0)	
10-50	15	14 (93.3)	1 (6.7)	
50-100	17	13 (76.5)	4 (23.5)	0.137
100-150	9	6 (66.7)	3 (33.3)	
>150	22	13 (59.1)	9 (40.9)	
Lymphocyte				
Normal	21	16 (76.2)	5 (23.8)	
Abnormal	43	31 (72.1)	12 (27.9)	1.000
Neutrophils				
Normal		22 (84.6)	4 (15.4)	
Abnormal		25 (65.8)	13 (34.2)	0.149

SD = standard deviation; *WBC* = white blood cell; *CRP* = *C*-reactive protein; eGFR = estimate glomerular filtration rate; GGO = ground glass opacity.

*Statistically significant.

Table 4 (cont'd.): Association between patients' clinical and laboratory findings and final outcome

Variable	Total	n (%)		<i>P</i> value
eGFR in mL/	min/1.73	m ²		
≥90	40	31 (77.5)	9 (22.5)	0.122
60-90	16	11 (68.8)	5 (31.3)	0.132
30-60	4	3 (75.0)	1 (25.0)	
15-30	2	0 (0)	2 (100)	
<15	2	2 (100)	0 (0)	
Consolidatio	n			
Yes	50	34 (68.0)	16 (32.0)	0.000
No	14	13 (92.9)	1 (7.1)	0.089
GGO				
Yes	58	42 (72.4)	16 (27.6)	1.000
No	6	5 (83.3)	1 (16.7)	1.000
Pleural effusi	ion			
Yes	2	1 (50.0)	1 (50.0)	0.464
No	62	46 (74.2)	16 (25.8)	0.464
Progression	pattern			
Initial deterio- ration followed by improve- ment (type I)	11	9 (81.8)	2 (18.2)	
Fluctuation (type II)	1	1 (100)	0 (0)	
Static (type III)	18	11 (61.1)	7 (38.9)	0.014*
Progressive deterio- ration (type IV)	12	5 (41.7)	7 (58.3)	
Progressive improve- ment (type V)	18	17 (94.1)	1 (5.9)	

SD = standard deviation; WBC = white blood cell; CRP = C-reactive protein; eGFR = estimate glomerular filtration rate; GGO = ground glass opacity.

*Statistically significant.

The median score of initial CXR for deceased patients was significantly higher than for those who recovered (17 versus 11; P = 0.009). In addition, there was a higher median score for initial CXRs for those who were admitted to the ICU than those who were not admitted to the ICU (14 versus 2.5; P = 0.0001). Similarly, higher median scores of the initial CXRs were observed for those who were intubated compared to those who were not intubated (16 versus 5; P = 0.0001) [Table 3].

 Table 5: Multivariate binary logistic regression analysis

 to determine the independent predictors of mortality

Variable	В	P value	OR (95% CI)		
Age	0.082	0.004*	1.085 (1.027-1.147		
Total baseline CXR score	0.107	0.044*	1.113 (1.003–1.236)		
Diabetes mellitus					
No (reference)					
Yes	0.936	0.207	2.550 (0.595-10.928)		
WBC count					
Normal (reference)					
Abnormal	1.420	0.065	4.138 (0.916-18.700)		

OR = odds ratio; CI = confidence interval; CXR = chest X-ray; WBC = white blood cell.

*Statistically significant.

A significant positive correlation was found between the duration of hospitalisation and the score of the initial CXR (correlation coefficient = 0.381; *P* = 0.002). There was no significant correlation between the initial CXR and duration of ICU admission or intubation.

Upon review of the follow-up CXRs, five evolution patterns were identified: type I (initial deterioration followed by improvement), type II (fluctuate), type III (static), type IV (progressive deterioration) and type V (progressive improvement). Type III and IV progression patterns were the most frequent among the deceased group. Age (P = 0.001), diabetes mellitus (P= 0.044) and WBC count (P = 0.034) were significantly associated with the final outcomes of the patients in the univariate analysis [Table 4].

There was a statistically significant difference in the mortality rate between the comorbidity groups (P = 0.004). The mortality rate among patients who had no comorbidity was 10.3% (n = 3), whereas the mortality rates among single comorbidity and multiple comorbidity groups were 56.3% (n = 9) and 26.3% (n = 5), respectively.

A multivariate binary logistic regression was performed to determine the effects of age, baseline CXR score, diabetes mellitus and WBC count on the likelihood of mortality among patients with COVID-19 pneumonia. The logistic regression model was statistically significant (χ^2 [4] = 24.130; *P* = 0.0001). The model explained 45.8% (Nagelkerke R²) of the variance in the outcome mortality and correctly classified 87.5% of cases. Increasing age (odds ration [OR] = 1.085, 95% confidence interval [CI]: 1.027–1.147; *P* = 0.004) and baseline CXR score (OR = 1.113,

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Figure 1: Chest X-rays (CXRs) showing the division of each lung divided into three equal zones with each zone given a score from 0 to 4 based on the percentage the lung involved. A: Initial frontal chest radiograph of 40-year-old man with COVID-19 showing clear lung filed (CXR score = 0). **B:** Initial frontal CXR of a 57-year-old male with COVID-19 showing right middle zone consolidation along with ground-glass opacities in left lower lung zone (CXR score = 2). **C:** Initial frontal CXR of a 64-year old male with COVID-19 showing right and left middle zones ground-glass opacity, right and left lower zones consolidation as well as ground-glass opacity (CXR score = 7). **D:** Follow-up frontal CXR of a 67-year old male with COVID-19 showing bilateral diffuse consolidation and ground-glass opacities (CXR score = 24). *RULZ = right upper lung zone; RMLZ = right middle lung zone; RLLZ = right lower lung zone; LULZ = left upper lung zone; LMLZ = left middle lung zone; LLZ = left lower lung zone; LLZ = left middle lung zone*

95% CI: 1.003–1.236; P = 0.044) were significantly associated with an increased likelihood of mortality [Table 5].

Discussion

A combination of clinical suspicion, RT-PCR and CXR and CT imaging is used to diagnose COVID-19.^{5,23} Although CXR is less sensitive when compared to CT, as well as due to the infection control issues and lack of availability of CT machines in many parts of the world, CXR is an alternative imaging modality that can be used for identification and follow-up of lung abnormalities in patients with COVID-19.^{11,24}

In the current study, GGO was the most common CXR manifestation of COVID-19 pneumonia (96.7%), followed by consolidation (83.3%). Mixed peripheral and central distribution of lung abnormalities was the most common distribution (60.0%) followed by peripheral distribution alone (33.3%). Predilection for lower lung zones involvement was another feature of COVID-19 pneumonia on CXR. Pleural effusion was rare as it was seen in only two patients. The absence of cavitation is an important negative finding that, when present, requires consideration of an alternative diagnosis or co-existing or added superinfection. the current study's results are consistent with the findings reported in several recent studies.^{11,12}

CXR findings of COVID-19 pneumonia can overlap with other viral pneumonias, in particular, other coronavirus infections, due to the similar pathogenesis of viruses belonging to the same family. For example, SARS and Middle East respiratory syndrome (MERS) share common CXR manifestations with COVID-19.¹⁷ Therefore, these findings need to be interpreted in combination with the clinical context of patients.

The CXR score was found to be an independent predictor of mortality and morbidity of admitted patients with COVID-19 pneumonia in the current study. A higher CXR score is associated with increased duration of hospitalisation, rate of ICU admission and intubation along with increased fatality. Borghesi et al. used a different scoring system to quantify the severity of CXR findings of patients with COVID-19 pneumonia. Each CXR was divided into three zones and each zone was given a score as follows: 0 was no lung abnormality, 1 was intestinal infiltrates, 2 was intestinal and alveolar infiltrates (interstitial predominance) and 3 was intestinal and alveolar infiltrates (alveolar predominance). Findings indicated that a high CXR score is associated with higher inhospital mortality.25,26

Regarding the temporal evaluation of lung abnormalities in the current study, progressive deterioration (type IV pattern) and static findings (type III pattern) were the most common evolution patterns among the deceased group, whereas the progressive improvement pattern was the most frequently observed pattern among recovered patients. Wong *et al.* studied the pattern of progression of 138 patients with SARS and found the most common pattern was initial radiographic deterioration to peak level followed by radiographic improvement.²⁷ Das *et al.* studied the progression pattern in 55 patients with MERS and found that progressive deterioration was the most common progressive pattern.²⁸ This study is an adjunct to the current literature since it proposes another quantifying method to assess the severity of CXR findings in patients with COVID-19 that can help stratify patients and plan their management.

Older age and diabetes mellitus were significantly associated with an increased likelihood of mortality. These findings are consistent with results from prior studies.²⁹

This study was subject to certain limitations. First, this study had a small sample size. Second, due to this study's retrospective nature, some important clinical information might not be well documented (for example, viral load, oxygen saturation and detailed symptoms at presentation). Lastly, visual estimation of the lung zones and percentage of lung involvement are subjective; therefore, the percentage of lung involvement might be over- or under-estimated.

Conclusion

A high score of the initial CXR in patients with COVID-19 pneumonia, accompanied by increased age, diabetes mellitus and increased WBC count, were associated with a poor prognosis and higher mortality. This information could help clinicians stratify and manage patients with COVID-19 pneumonia better.

CONFLICT OF INTEREST

The authors declare no conflicts of interest.

FUNDING

No funding was received for this study.

AUTHORS' CONTRIBUTION

RSU and MAB collected the data. RSU, IAS and JAK reviewed the chest X-ray. RSU drafted the manuscript. SJ performed the statistical analysis and wrote the statistical section. AK, SAT, FAK, FK and HAK reviewed the manuscript and provided intellectual input. All authors approved the final version of the manuscript.

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