

## RESEARCH / INVESTIGACIÓN

## Clinical Characteristics and Outcome of SARS-CoV-2 Patients. An Experience from Anbar province - West of Iraq

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**Abstract:** The SARS-CoV2 infection emerged in Iraq in February 2020. In this study, we describe the clinical characteristics and outcomes of the initial SARS-CoV2 patients. A total of 529 patients were included in this study from April to August 2020 in Anbar province. Patients were confirmed to be infected in nasal swabs by real-time RT-PCR or chest CT scan findings. The gathered data included the demographic variables (age, sex, residency), presence of comorbidity (hypertension, diabetes mellitus, respiratory illness, coronary heart disease, chronic kidney disease, obesity), and history of contact with a known case of SARS-CoV2. The results showed that 64% of the patients were males and 36% were female, 48% of the patients lied in the age category 40-59 years, 74% had exposure history, 95% did not have a history of smoking, 46% were overweight, 60% had no comorbidity, 78% presented with mild/moderate disease, 70% had typical chest CT scan finding (CO-RAD 5), and 76% of patients showed positive PCR. The fatality rate is 16%. Most of the patients had a history of exposure to a confirmed case of SARS-CoV2 before the illness. The severity and outcome were correlated with risk factors and comorbidity. Combining chest computed tomography images with the qPCR analysis of nasal swab samples can improve the accuracy of SARS-CoV2 diagnosis.

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**Key words:** SARS-CoV-2, Clinical Characteristics, Outcome, West of Iraq.

## Introduction

SARS-CoV2 is a novel virus causing a global pandemic with significant morbidity and mortality. It is a coronavirus from Coronaviridae's family and was first discovered in Wuhan, China, in December 2019; it is also named Covid-19 (Corona Virus Disease 2019)<sup>1</sup>. The discovery of the virus was made after having an increasing number of patients with acute respiratory tract infection not attributed to known viral infections like the influenza virus, avian flu, and severe acute respiratory syndrome coronavirus (SARS-CoV). Lockdown of the area where the disease appeared first with other precautionary measures has failed to contain the virus, and it spread rapidly to other countries in Asia (South Korea, Iran, India, Pakistan, and many others) and then globally<sup>2,3</sup>. In its update in March 2020, the World Health Organization (WHO) declared the disease a pandemic in a press announcement. This is not the first outbreak caused by this kind of viruses in the last 2 decades, since it had been described in 2002 (Severe Acute Respiratory Syndrome Coronavirus (SARS-CoV) and 2012 (the Middle East Respiratory Syndrome Coronavirus (MERS-CoV)). The mortality rate of SARS-CoV was 10%, while that of MERS-CoV was 37%<sup>4,5</sup>. The spread of the virus varied enormously among different countries and within the same country or region. However, many studies have confirmed that the number of affected cases tremendously overshoots reported cases confirmed by PCR testing<sup>6</sup>.

The average incubation period of the disease is 6 days<sup>7</sup>, but it could be as long as 14 days<sup>8</sup>. Most patients develop only mild symptoms of fever, mild cough, loss of taste and smell sensations, and muscle cramps. Nonetheless, a minority might present with severe pneumonia, acute respiratory distress syndrome, and multi-organ failure<sup>9,10</sup>. Radiologically, bilateral pulmonary parenchymal ground-glass and consolidative pulmonary opacities are the most frequent CT; however, occasionally, rounded morphology and peripheral lung distribution might be evident<sup>11,12</sup>. Iraq reported the first case of covid-19 in

late February 2020 in Al-Najaf province. The disease then gradually involved all governorates and provinces. The first confirmed case in Anbar Province (West of Iraq) was on 25th March 2020, and the first death was on 31st May 2020<sup>13</sup>. Knowledge of the natural course of the disease and its virulence is essential in organizing proper plans and procedures to face this illness effectively. Studies have shown that Covid-19 has a wide range of disease severity and mortality<sup>14</sup>, representing a fundamental challenge facing health care providers. Several studies were carried out in different parts of Iraq targeting these objectives, and this study represents another one to describe the clinical characteristic and outcomes of the disease in the West of Iraq.

## Materials and methods

Depending on the local protocol, patients who tested positive to SARS-CoV2 using Real-Time Polymerase Chain Reaction (RT-PCR) and patients with classical clinical and radiological features and history of contact with Covid-19 patients (even with negative RT-PCR) were enrolled in this cross-sectional study. We used a CT-based system (CO-RADS ) to assess the suspicion of pulmonary involvement in SARS-CoV2<sup>15</sup>. Patients were either admitted to Ramadi Teaching Hospital or treated at home based on their clinical status. Those who were treated at home were followed by medical personnel regularly. The management line is based on the Iraqi Ministry of Health's national guidelines for clinical care and treatment of SARS-CoV2, updated regularly.

A total of 529 patients were included in this study during the period from April to August 2020. The gathered data included the demographic variables (age, sex, place of living), presence of comorbidity (Hypertension, diabetes mellitus, respiratory illness, coronary heart disease, chronic kidney disease,

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se), and history of contact with a known case of SARS-CoV2. Patients were categorized into 4 categories. Mild disease is when a patient complains of one or more of the following: Low-grade fever, mild cough, runny nose, and sore throat. The moderate disease is when a patient shows clinical and radiological features of pneumonia without hypoxia: Fever, cough, chest tightness, clinical and radiological evidence of consolidation(s). Severe disease is when a patient shows clinical and radiological features of pneumonia with hypoxia which requires assisted ventilation. Acute disease is when a patient suffers from clinical shock or organ(s) failure<sup>16</sup>. The Human Research and Ethics Committee, Anbar University (Ministry of Higher Education and Scientific Research- Iraq) approved this study (Notification letter No. 69/2020).

### Statistical analysis

It was carried out using SPSS, version 22. Since all variables were of categorical types, the Chi-Square test with a 5% significance level was used to examine the investigated relationships between patients' characteristics and survival status. If the chi-square test's assumption (expected count) was violated, either Fisher's exact test or likelihood ratio Chi-square test was used as suitable.

### Results

During the study period, 529 patients with SARS COV-2 were either admitted in Ramadi hospital or followed at home. Of them, 64% were males, 48% of age 40-59 years, 74% with exposure history, 95% without a history of smoking, 46% overweight, 60% without comorbidity, 78% with mild/moderate covid, 70% having CO-RAD 5 chest CT scan finding and 76% with positive PCR. Out of 529 covid-19 patients, 84% survived, while the remaining 16% could not survive (case mortality rate is 16%).

Table (1) shows the characteristics of SARS-CoV2 patients by survival status. There was no significant difference in the distribution of survival status by gender. The percentage of non-survival increased with age, and the association was significant ( $p=0.001$ ). The majority of non-survivors had no smoking history. The highest percentage of non-survivors was among overweight patients (35%), and the difference was statistically significant ( $p=0.001$ ). Almost all non-survivors (99%) had at least one comorbidity (hypertension, diabetes, coronary heart disease, chronic obstructive lung disease, chronic kidney disease) compared to 29% of survivors, and the association was significant ( $p=0.001$ ).

Table (2) survivors reported less exposure history (72%) compared to non-survivors (86%), and the difference was statistically significant ( $p=0.008$ ). All non-survivors (100%) were diagnosed with severe/critical covid-19 status compared to only 8% of survivors, and the association was significant ( $p=0.001$ ).

The percentage of non-survivors increased with disease severity, and CORAD level as shown by CT scan as 85% of non-survivors had CO-RAD 5 ( $p=0.001$ ) (Table 3).

### Discussion

SARS-CoV2 pandemic continues spreading despite all measures exerted by health authorities to face it. Understanding patients' characteristics about the survival rates of the disease is valuable in this confrontation. We are still gaining knowledge in identifying clinical characteristics of severe disease and mortality. This study summarizes the clinical characteristics of 529 cases with SARS-CoV2 infection concerning the outcome, i.e., survivors vs. non-survivors. Knowing these characteristics will have a significant impact on improving survival. The reported overall case fatality in different Iraq provinces is as follows: Southern 9.4%, Northern 2.1%, Eastern 16.7%, and the Capital 8.7%<sup>17</sup>. This study estimated 16% as a case fatality rate in Anbar province – west of Iraq.

There is a consensus among researchers and clinicians that the increasing age of the patients is significantly linked to a higher non-survival rate in SARS-CoV2 patients<sup>18,19</sup>. Likewise, the current results showed that elderly patients were prone to more severe disease and higher mortality. The possible cause is not well understood; however, viable hypotheses emerge, including changes to the immune cell repertoire, epigenome, nicotinamide adenine dinucleotide (NAD+) levels, and inflammation activity biological clocks, and covalent modifications of human and viral proteins<sup>20</sup>.

Several studies confirmed that SARS-CoV2 infection is more common in patients with comorbidities<sup>1,10,18</sup>. This is consistent with the current study. However, the results could not find any dominance of particular comorbidity over the others in terms of outcome. Factors that could explain this include the increased expression of the angiotensin-converting enzyme-2 (ACE2), cytokine storm, and drug interactions in patients with comorbidities<sup>21</sup>.

Many studies have mentioned that the male gender is one of the risk factors for increased severity and mortality independent of age<sup>22-24</sup>. However, this study showed no significant difference in terms of survival status and gender. Similar fin-

| Demographic |             | Survival Status  |     |                     |     | p-value |               |
|-------------|-------------|------------------|-----|---------------------|-----|---------|---------------|
|             |             | Survivor (n=444) |     | Not Survivor (n=85) |     |         | Total (n=529) |
| sex         | Male        | 282              | 64% | 56                  | 66% | 338     | 0.677         |
|             | Female      | 162              | 36% | 29                  | 34% | 191     |               |
| Age group   | <25 years   | 22               | 5%  | 1                   | 1%  | 23      | 0.001         |
|             | 25-39 years | 106              | 24% | 11                  | 13% | 117     |               |
|             | 40-59       | 229              | 52% | 25                  | 29% | 254     |               |
|             | >=60 years  | 87               | 20% | 48                  | 56% | 135     |               |

**Table 1.** Survival status of SARS-CoV2 patients by demographic characteristics.

| Risk Factors     |  | Survival Status  |     |                     |     |               | p-value |
|------------------|--|------------------|-----|---------------------|-----|---------------|---------|
|                  |  | Survivor (n=444) |     | Not Survivor (n=85) |     | Total (n=529) |         |
| Smoking          | Yes  | 24               | 5%  | 4                   | 5%  | 28            | 0.522   |
|                  | No   | 420              | 95% | 81                  | 95% | 501           |         |
| Body weight/ BMI | Normal (18.5-24.9)   | 178              | 40% | 26                  | 31% | 204           | 0.001   |
|                  | Over weight (25-29.9)  | 215              | 48% | 30                  | 35% | 245           |         |
|                  | Obesity (30-34.9)  | 36               | 8%  | 20                  | 24% | 56            |         |
|                  | Morbid Obesity (+40)   | 15               | 3%  | 9                   | 11% | 24            |         |
| Comorbidity      | Yes  | 127              | 29% | 84                  | 99% | 211           | 0.001   |
|                  | No   | 317              | 71% | 1                   | 1%  | 318           |         |
| Comorbidity      | Hypertension   | 31               | 7%  | 45                  | 53% | 76            | 0.001   |
|                  | Diabetes   | 21               | 5%  | 55                  | 65% | 76            | 0.001   |
|                  | Coronary heart disease   | 14               | 3%  | 12                  | 14% | 26            | 0.001   |
|                  | Chronic obstructive lung disease   | 8                | 2%  | 11                  | 13% | 19            | 0.001   |
|                  | Chronic kidney disease   | 4                | 1%  | 2                   | 2%  | 6             | 0.248   |
|                  | Others(Stroke, Chronic liver diseases, Rheumatoid Arthritis and Malignancies ) | 6                | 1%  | 2                   | 2%  | 8             | 0.621   |

**Table 2.** Survival status of SARS-CoV2 patients by risk factors assessment.

| COVID Status     |                 | Survival Status  |     |                     |      |               | p-value |
|------------------|-----------------|------------------|-----|---------------------|------|---------------|---------|
|                  |                 | Survivor (n=444) |     | Not Survivor (n=85) |      | Total (n=529) |         |
| Exposure history | Yes             | 321              | 72% | 73                  | 86%  | 394           | 0.008   |
|                  | No              | 123              | 28% | 12                  | 14%  | 135           |         |
| Disease severity | Mild/Moderate   | 410              | 92% | 0                   | 0%   | 410           | 0.001   |
|                  | Severe/Critical | 34               | 8%  | 85                  | 100% | 119           |         |
| Chest CT-Scan    | CORAD3          | 21               | 6%  | 0                   | 0%   | 21            | 0.001   |
|                  | CORAD4          | 88               | 27% | 12                  | 16%  | 100           |         |
|                  | CORAD 5         | 217              | 67% | 65                  | 84%  | 282           |         |
| PCR              | POSTIVE         | 327              | 75% | 70                  | 82%  | 397           | 0.129   |
|                  | NEGATIVE        | 111              | 25% | 15                  | 18%  | 126           |         |

**Table 3.** Survival status of SARS-CoV2 patients by disease exposure and severity.

dings have been seen in other studies<sup>25-27</sup>. A possible explanation would be that in certain countries, like Iraq, women are at higher risks due to demographic factors or local health characteristics which has been shown in other studies conducted in other parts of the country<sup>17,28</sup>. The history of smoking was considered as a nasty prognostic factor in numerous studies<sup>1,29-32</sup>. This is in contrast with the current findings, as the majority of non-survivors were a non-smoker. This may be related to the low numbers of smokers in this study compared to non-smokers or incomplete records. In addition to this, severe SARS-CoV2 targets the older population (>65 years), in whom smoking rates are approximately 3-5 fold lower than that in the general population<sup>33</sup>. Obese patients are at higher risk of developing severe and critical illness than non-obese patients<sup>34-36</sup>. A similar finding was shown in this study. This is likely because obese patients are known to have a defective immune system in addition to underlying comorbidities<sup>37,38</sup>. Obesity changes the role of immunity by altering the response of cytokines, resulting in a decrease in the cytotoxic cell response of immuno-competent cells that have a key anti-viral role and disturb the balance of endocrine hormones, like leptin, that affect the interaction between metabolic and immune systems<sup>39,40</sup>. Chest CT scan is a valuable tool in the diagnostic process of viral pneumonia cases associated with SARS-CoV2. The sensitivity and specificity of the chest CT in diagnosing SARS-CoV2 and the radiation exposure have to be judged together. Arguments exist regarding the value of chest CT scan for SARS-CoV2 diagnosis, particularly those patients who exhibit typical clinical symptoms and have negative RT-PCR results in highly infected regions<sup>41</sup>.

## Conclusions

The SARS-CoV2 infection caused severe respiratory illness with significant morbidity and mortality. Understanding these factors can enhance defining patients at higher risk and allow a more targeted approach to prevent those deaths. Using chest CT scan images with nasal swab sample qPCR analysis may improve the accuracy of SARS-CoV2 diagnosis.

## Conflict of interest

Authors have no conflict of interest.

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