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# The Impact of SARS-CoV-2 on the Mental Health of Patients with Primary Immune Deficiency and Their Parents

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#### **Abstract**

Since the start of the Coronavirus disease (COVID-19) pandemic, people have had to cope with significant mental pressure regarding this mysterious infection. The rapid spread of syndrome coronavirus 2 (SARS-CoV-2) worldwide has led to immense anxiety and depression among people, especially in the immunocompromised population and their close relatives. This study aims to evaluate the mental health of patients with primary immune deficiency (PID) and their mothers during the first year of the COVID-19 outbreak. A cross-sectional study was conducted from September 2020 to March 2021 among pediatric patients with PID and their family members. Thirty PID patients and their mothers were investigated using the Zung Self-Rating Anxiety Scale and Beck Anxiety Inventory (BAI), respectively. The results showed that mothers of younger PID patients experienced higher levels of anxiety (P-value = 0.019). Additionally, PID patients who did not comply with quarantine measures had significantly higher anxiety levels (P-value = 0.02). The results indicate mild to moderate anxiety in PID patients aged  $12.1 \pm 1.46$  years during the first year of the COVID-19 outbreak. Furthermore, there was mild anxiety observed in mothers of PID patients. Non-compliance with quarantine measures significantly contributed to higher anxiety scores in both groups, emphasizing the importance of staying home and adhering to restricted social interactions to improve mental health.

Keywords: Primary Immune Deficiency, COVID-19, SARS-Cov-2, Anxiety

## 1. Background

Since the emergence of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) in 2019, a wide range of disease manifestations and complications has been observed. While respiratory symptoms are among the most prevalent presentations, other organs might be involved in different ways (1, 2). That's why complications that are expected as a consequence of this infection are very different.

The World Health Organization (WHO) introduced SARS-CoV-2 as a major threat to mental health (3). People worldwide experienced a variety of psychological responses during the Coronavirus disease (COVID-19) outbreak. When people faced an unknown viral

infection with a breakneck speed of spread, they normally responded to that with fear, anxiety, and even depression. These, alongside the economic burden induced by the pandemic, affect people in different aspects (4). Moreover, stay-at-home protocols of governments in infection prevention can significantly worsen the situation. Lack of physical contact with family and friends, especially when they are experiencing such tough moments, could worsen the mental problems (5, 6).

Eventually, WHO and other relevant organizations provided multiple guidelines for improving mental health and screening high-risk people such as children, older adults, and those with chronic mental illnesses, among others. Anxiety is one of the psychological

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problems that can be accompanied by fear, excessive concern, and mood instability. One of these high-risk populations comprises people who have underlying susceptibility to infections like primary immune deficiencies (PIDs) (7). In the first months of the SARS-CoV-2 outbreak, many of our PID patients stopped visiting hospitals. They preferred online visits, which caused multiple difficulties in managing their problems on time (8). So, we conducted this study to examine their psychological status over time.

#### 2. Methods

A cross-sectional study was conducted from September 2020 to March 2021 among the pediatric population with PID and their family members. Psychological assessment in children was based on the Zung Self-Rating Anxiety Scale, and we also evaluated anxiety in their mothers using the Beck Anxiety Inventory (BAI). Both of their Persian formats were validated by previous studies (9). The Zung Scale includes 20 multiple-choice questions (10). Each question is scored between 0 and 3, so the final score will range from 0 to 60. Scores above 26 are considered severe anxiety, while scores between 8 and 15 and between 16 and 25 are considered mild and moderate anxiety, respectively.

In the BAI, there are 21 multiple-choice questions describing common symptoms of this psychological problem, including mental, physical, and panic symptoms. Scoring is the same as the Zung Scale, but the final score will be 63. Each question is scored between 0 and 3. Scores above 26 are considered severe anxiety, while scores between 8 and 15 and between 16 and 25 are considered mild and moderate anxiety, respectively. After obtaining consent from the parents of the children, they were evaluated by a psychologist. Data were analyzed using SPSS 22, and the Shapiro-Wilk and Kolmogorov-Smirnov tests were performed to check the normal distribution of the quantitative variables of age and the total score of the questionnaires. Ethical code: IR.MUMS.REC.1399.155.

### 3. Results

Among 30 patients with PID, with a mean age of  $12.1 \pm 1.4$  years (range: 4 - 21 years), humoral immune

deficiency was the most prevalent (30%). Other types of PIDs in our participants are listed in Table 1. Approximately eighteen participants (60%) had to regularly admit to the hospital and receive intravenous immunoglobulin (IVIG). Others did not need IVIG at all. There was an equal frequency of compliance with quarantine. During the study period, 16 patients (53.3%) experienced at least one symptom attributed to infection, such as fever, chills, cough, fatigue, or sneezing. They underwent nasal PCR for SARS-CoV-2; however, only 5 patients had positive results.

The mean score of the Zung Self-Rating Anxiety Scale in primary immune deficient patients was  $16.4 \pm 4.2$  (range: 9 - 26). There wasn't any statistically significant relationship between the total Zung score and age (P-value = 0.876) or sex (P-value = 0.98). However, there was a significant relationship between anxiety and regular hospitalization for IVIG administration (P-value = 0.001). In addition, those PID patients who didn't comply with quarantine had significantly more anxiety (P-value = 0.02). More details are in Table 2.

The mean score of the BAI in mothers of PID patients was  $8.56 \pm 7.45$  (range: 0 - 22). Because the distribution of this score was abnormal, we analyzed it using Spearman's statistical analysis. There was a significant relationship between the age of children and the anxiety of their mothers. As a result, mothers of younger PID patients had more anxiety (P-value = 0.019). In addition, there was more anxiety in those mothers whose children experienced suspected COVID-19 manifestations and those whose families didn't comply with quarantine. However, regular hospitalization for IVIG did not cause a higher anxiety score in mothers. More details are in Table 3.

#### 4. Discussion

After the emergence of SARS-CoV-2, strict regulations were put in place to enforce staying at home, which restricted social interactions and led to major concerns about this mysterious contagious infection. There wasn't any effective approved preventive or treatment strategy, so mental health was treated as a major concern, especially in immunocompromised patients (3, 5, 6). We conducted this study in the first year of the COVID-19 outbreak, aiming to investigate anxiety in

Table 1. Demographic Characteristics of Primary Immune Deficiency Patients Participating in Our Study a

Variables	Frequency
Sex (male/female)	17/13
Type of PID	
Humoral immune deficiency (CVID, Broton)	9 (30)
CID	2 (6.7)
Phagocytic disorder (neutropenia, CGD, LAD)	5 (16.7)
MSMD	2 (6.7)
Syndromic disease (hyper IgE, ataxia telangiectasia, wiskott aldrich)	12 (40)

Abbreviation: PID, primary immune deficiency.

<sup>a</sup>Values are expressed as No. (%).

Table 2. Mean Scores of the Zung Self-rating Anxiety Scale		
Zung Self-rating Anxiety Scale	$\textbf{Mean} \pm \textbf{Standard Deviation}$	P-Value <sup>a</sup>
Total	$16.4 \pm 4.2$	
Female	$16.38 \pm 3.7$	0.98
Male	$16.41 \pm 4.8$	
People which had regular visit at hospital for IVIG infusion	$19.50 \pm 4.18$	0.001
People who didn't need regular hospitalization for IVIG	$14.33 \pm 2.97$	
Comply with quarantine	14.94 ± 2.55	0.02
Didn't comply with quarantine	18.57 ± 5.46	
Experience of respiratory infection manifestation during study	16.1 ± 3.72	0.59
Didn't have any symptoms attributing to the respiratory infection	$16.85 \pm 4.97$	

 $^{\mathrm{a}}$  Independent t-test.

children with inborn errors of immunity and their mothers. According to our data, there was mild to moderate anxiety in PID patients aged  $12.1 \pm 1.46$  years during the first year of the COVID-19 outbreak. Additionally, there was mild anxiety in mothers of PID patients. These results are in agreement with Akdag et al., who evaluated anxiety in parents of PID patients during the COVID-19 outbreak and found that it was significantly higher than in parents of healthy children (11). Also, in the study by Sowers and Galantino, anxiety was higher in the pediatric population with PID during the outbreak compared to before the outbreak (12).

In a study conducted by Topal et al. in 2020, anxiety was evaluated in parents of children with PID who had to receive IVIG monthly. They found that these parents had a higher anxiety score than parents of non-PID children (P = 0.003), which contrasts with our study (13). In our study, all patients had inborn errors of immunity, so we expected parents to be concerned about infection

in their immunocompromised child. Therefore, regular hospitalization or non-hospitalization did not lead to a significant difference between the two groups.

One of our limitations was the non-availability of anxiety data in the general population in Iran, especially during the outbreak. This emphasizes the need for larger studies conducted in healthy people. Additionally, follow-up studies in immunocompromised patients after the COVID-19 outbreak will be helpful. Comparing the results will illustrate the impact of COVID-19 on the mental health of these susceptible populations.

## 4.1. Conclusions

According to our data, there was mild to moderate anxiety in PID children and mild anxiety in their mothers during the first year of the COVID-19 outbreak. Non-compliance with quarantine significantly led to higher anxiety scores in both groups. Therefore, advice

Table 3. Beck Anxiety Inventory in Mothers of PID Patients Mean ± Standard Deviation **Beck Anxiety Inventory** P-Value <sup>a</sup>  $8.56 \pm 7.45$ People which had regular visit at hospital for IVIG infusion 14.81 ± 1.2 0.59 People who didn't need regular hospitalization for IVIG 16.54 ± 2 Comply with quarantine 11.56 ± 1.5 0.03 Didn't comply with quarantine  $21.42 \pm 3.2$ Experience of respiratory infection manifestation during study 18.56 ± 2.5 0.04 Didn't have any symptoms attributing to the respiratory infection  $12.2 \pm 1.3$ 

Abbreviation: IVIG. intravenous immunoglobulin.

to stay home and more restricted social communication improved mental health.

#### **Footnotes**

Authors' Contribution: M.B.K. and H.A. conceived and designed the evaluation, and drafted the manuscript. M.B.K. participated in designing the evaluation, performed parts of the statistical analysis, and assisted in drafting the manuscript. N.M., M.M., A.A., and H.A. reevaluated the clinical data, revised the manuscript, conducted the statistical analysis, and further revised the manuscript. M.B.K., A.A., and H.A. collected the clinical data, interpreted them, and revised the manuscript. M.B.K., N.M., and H.A. re-analyzed the clinical and statistical data and revised the manuscript. All authors read and approved the final manuscript.

**Conflict of Interests Statement:** The authors declare no conflicts of interest.

**Data Availability:** The dataset presented in the study is available on request from the corresponding author during submission or after publication.

**Ethical Approval:** Our study protocol has been reviewed and approved by the Health Research Ethical Committee of the Faculty of Medicine, Mashhad University of Medical Sciences. The ethical code is IR.MUMS.REC.1399.155.

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**Informed Consent:** Informed consent was obtained from the parents of the children.

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<sup>&</sup>lt;sup>a</sup> Mann-Whitney test.

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