Reinfection with SARS-CoV-2 after 152 Days: A Case Report
SM Abdullah Al Mamun, Quazi Tarikul Islam, Sanaullah Sarker, Asifur Rahman

Abstract

Background: In general, reinfection means a person was infected (got sick) once, recovered, and then later became infected again. Based on what we know from similar viruses, some reinfections are expected. We are still learning more about COVID-19. Ongoing COVID-19 studies will help us to understand. The degree of protective immunity conferred by infection with severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) is currently unknown. As such, the possibility of reinfection with SARS-CoV-2 is not well understood. Here we describe presentation and investigations of two instances of SARS-CoV-2 infection in the same individual.

Methodology: A 38-year-old man who is a resident of Dhaka city, Engineer by profession presented to doctors on two occasions with symptoms of viral infection. First time at a community testing event in June, 2020, and a second time to Evercare hospital at the mid November, 2020. Nasopharyngeal swabs were obtained from the patient at each presentation and twice during follow-up. RT PCR testing was done to confirm SARS-CoV-2 infection. CXR P/A view was performed in 1st time of Covid infection & HRCT of chest was performed during his stay at Evercare hospitals.

Findings: The patient had two positive tests for SARS-CoV-2, the first on June 20th, 2020, and the second on November 18th, 2020, separated by two negative tests done during follow-up in July, 2020. The second infection was symptomatically more severe than the first with remarkable radiological changes of Chest in high resolution CT scanning.

Interpretation: Two subsequent negative RT PCR after 1st positive with clinically asymptomatic period after 2 weeks of mild symptoms was observed in our patient. About 5 months later he was Covid19 positive again with moderate symptoms. These findings suggest that the patient was infected by SARS-CoV-2 on two separate occasions probably by genetically distinct virus. Thus, previous exposure to SARS-CoV-2 might not guarantee total immunity in all cases. All individuals, whether previously diagnosed with COVID-19 or not, should take identical precautions to avoid infection with SARS-CoV-2. The implications of reinfections could be relevant for vaccine development and application.

Key words: COVID-19, Reinfection, RT-PCR

Introduction
Infection with severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) leads to a detectable immune response, but the susceptibility of previously infected individuals to reinfection with SARS-CoV-2 is not well understood. SARS-CoV-2 infection results in generation of neutralizing antibodies in patients. However, the degree to which this immune response indicates a protective immunity to subsequent infection with SARS-CoV-2 has not yet been elucidated. Reports of secondary infection events with SARS-CoV-2 have been published from Hong Kong, the Netherlands and Belgium and Ecuador.

We present a case report of an individual who had two distinct COVID-19 illnesses on 5 months apart with more severity on 2nd episode.

Case history
We present a case report of a 38-year-old male patient who was a resident of Dhaka City. The patient presented to a...
community testing event held by the IEDCR on June 20, 2020 & was tested positive for COVID-19. He had symptoms consistent with viral infection (sore throat, cough, headache, nausea) which had started on June 15, 2020 (figure 1). The patient had no history of clinically significant underlying conditions, and no indications of compromised immunity were identified. During isolation at home, the patient’s symptoms resolved (reported on June 29, 2020) and he continued to feel well. His repeat Nasopharyngeal sample tested negative on two occasions 7 days apart. Until November 15, 2020 he was alright & full active. He performed many field activities as his official assignment & lastly he stayed at home for 5 days. From where he developed symptoms. On November 17, 2020, the patient sought care at ED of Evercare hospitals Dhaka with self-reported fever, headache, dizziness, cough, nausea for 2 days, at which time chest radiography was done and shows bilateral peripheral GGO with altered lab parameter (CRP, D-dimer, LDH, Ferritin, CBC) & he was admitted in Grey Zone of Evercare Hospitals according to Hospitals protocol. Next morning his Nasopharyngeal swab had been sent for SARS COV-2 RT PCR & after 3 hours report obtained. He was Positive for COVID-19. As the patient complaining shortness of breathing and SpO2 had been shown lower side (< 94%) in room air, supplemental O2 was started. Subsequently HRCT of Chest was performed and shows diffuse involvement of both lung.

**Procedures** Specimens were obtained from the patient by nasopharyngeal swab at the community testing event in 1st time, when the result was positive for RT-PCR & subsequent two sample during the period of isolation and recovery. Swabs were transported to the respective Laboratory on cold packs and stored by refrigeration (4–8°C) before real-time RT-PCR. Others Lab including CRP, D-dimer, LDH, Ferritin & CXR P/A view done to know the course of COVID-19 disease.

During the 2nd episode of disease, when the patient had been admitted to our hospitals for progressive worsening of symptoms, nasopharyngeal swab was collected from bedside by our trained technician & transported to our laboratory on cold packs and stored by refrigeration (4–8°C) before real-time RT-PCR as before. Subsequent sample during the period of admission was sent to the PCR Lab maintaining cold packs. Lab test including CRP, D-dimer, LDH, Ferritin & HRCT of chest view done to know the course of COVID-19 disease.

**Results**

The first nasopharyngeal swab, obtained at the community screening event on April 18, 2020, was positive for SARS-CoV-2 on real-time RT-PCR testing. Two subsequent nucleic acid amplification tests obtained after resolution of symptoms were negative for SARS-CoV-2 RNA (table 1). The patient’s symptoms returned on November 15, 2020 from community contact and he was admitted to hospital on November 17, 2020, at which time a Nasopharyngeal swab was obtained and was positive for SARS-CoV-2 infection by real-time RT-PCR testing. The patient required ongoing oxygen support in hospital and reported symptoms that included myalgia, cough, and shortness of breath. Chest radiography showed development of patchy, bilateral, interstitial opacities suggestive of viral or atypical pneumonia. With two episodes of symptoms consistent with COVID-19, and two specimens positive for SARS-CoV-2 separated by a period of 152 days.

Table 1: *Time line of symptom onset, molecular diagnosis*.

<table>
<thead>
<tr>
<th>Symptom onset</th>
<th>Positive real time RT-PCR</th>
<th>Symptom resolution</th>
<th>Negative real time RT-PCR</th>
<th>Negative real time for 2nd time</th>
<th>Symptom onset for 2nd time</th>
<th>Positive real time RT-PCR</th>
<th>Positive real time RT-PCR</th>
</tr>
</thead>
<tbody>
<tr>
<td>June 15&lt;sup&gt;th&lt;/sup&gt;</td>
<td>June 18&lt;sup&gt;th&lt;/sup&gt;</td>
<td>June 29&lt;sup&gt;th&lt;/sup&gt;</td>
<td>6&lt;sup&gt;th&lt;/sup&gt; July</td>
<td>16&lt;sup&gt;th&lt;/sup&gt; July</td>
<td>15&lt;sup&gt;th&lt;/sup&gt; November</td>
<td>18&lt;sup&gt;th&lt;/sup&gt; November</td>
<td>6&lt;sup&gt;th&lt;/sup&gt; December</td>
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</tbody>
</table>

Table-II: *Summary of laboratory results*

<table>
<thead>
<tr>
<th>Test Methodology</th>
<th>1&lt;sup&gt;st&lt;/sup&gt; Episode</th>
<th>2&lt;sup&gt;nd&lt;/sup&gt; Episode</th>
</tr>
</thead>
<tbody>
<tr>
<td>Real time RT-PCR</td>
<td>HRCT of Chest after 2 weeks</td>
<td>Real time RT-PCR after 2 weeks</td>
</tr>
<tr>
<td>Real time RT-PCR</td>
<td>HRCT of Chest</td>
<td>Real time RT-PCR after 2 weeks</td>
</tr>
<tr>
<td>Real time RT-PCR</td>
<td>HRCT of Chest after 3 weeks</td>
<td></td>
</tr>
<tr>
<td>Test Result</td>
<td>Positive</td>
<td>Peripheral GGO</td>
</tr>
<tr>
<td>Diffuse GGO</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Positive</td>
<td>Positive</td>
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<tr>
<td>Positive</td>
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Discussion:
Our case report presents details of the individual in Bangladesh to have symptomatic reinfection with SARS-CoV-2 after 152 days of first infection. Similar to observations with the reinfection case in Ecuador⁶, our patient showed increased symptom severity in their second infection, whereas the cases from Belgium and the Netherlands⁷ and Hong Kong⁸ did not show a difference in severity of symptoms. The mechanisms that could account for a more severe secondary infection can only be speculated. First, a very high dose of virus might have led to the second instance of infection and induced more severe disease.⁹ Second, it is possible that reinfection was caused by a version of the virus that was more virulent, or more virulent in this patient’s context. Third, a mechanism of antibody dependent enhancement might be the cause, a means by which specific Fc-bearing immune cells become infected with virus by binding to specific antibodies. This mechanism has been seen...
previously with the beta coronavirus causing severe acute respiratory syndrome.\textsuperscript{10} In that case, the patient recovered and was discharged from hospital.

The individual associated with these two SARS-CoV-2 infections had no immunological disorders that may influence reinfection. He was not taking any immunosuppressive drugs. The individual was negative for HIV by antibody and RNA testing (data not shown) and had no obvious cell count abnormalities.

It is possible that we have reported a case of continuous infection entailing deactivation and reactivation. However, for such a hypothesis to be true, a mutational rate of SARS-CoV-2 would be required that has not yet been recorded.\textsuperscript{11,12} Limitation in our case was genome sequence was not possible.

Our findings have implications for the role of vaccination in response to COVID-19. If we have truly reported a case of reinfection, initial exposure to SARS-CoV-2 might not result in a level of immunity that is 100% protective for all individuals. With respect to vaccination, this understanding is established, with influenza regularly showing the challenges of effective vaccine design.\textsuperscript{13} A major limitation of our case study is that we were unable to undertake any assessment of the immune response to the first episode of SARS-CoV-2.

\textbf{Conclusion:} Reinfection in COVID-19 is not a new issue now. One should keep eye on second infection where the features may be worse than first attack.

\textbf{References:}
\begin{enumerate}
  \item Prado-Vivar B, Becerra-Wong M, Guadalupe JJ, et al. COVID-19 re-infection by a phylogenetically distinct SARS-CoV-2 variant, first confirmed event in South America. SSRN 2020;
  \item Mercatelli D, Giorgi FM. Geographic and genomic distribution of SARS-CoV-2 mutations. Front Microbiol 2020; 11: 1800
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