Positive Pleural Fluid RT-PCR for Virus Detection in SARS-CoV-2 Pneumonia

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Learning points for clinicians

This case report highlights that SARS-CoV-2 can be detected in pleural effusions by RT-PCR. In this case, the SARS-CoV-2 related pneumonia was associated with a lymphocytic, exudative, complicated pleural effusion.

Patient Presentation:

A 71-year-old African American man presented with a three-week history of fever, productive cough and progressively worsening shortness of breath to the emergency department. A RT-PCR COVID-19 test (Xpert Xpress SARS-CoV-2, Cepheid, Sunnyvale, CA) was performed via nasopharyngeal swab on arrival to the ED, and the patient was found to be positive. The patient had a rapidly deteriorating course with increased work of breathing and oxygen requirement and, therefore, was transferred to the intensive care unit. On presentation to the intensive care unit, his physical exam showed: Temperature 102.6 °F; heart rate 92 beats per minute; respiratory rate of 30 breaths per minute; blood pressure 157/95 mm Hg; and SpO2 of 95% on 2L nasal cannula. He was in mild respiratory distress. Chest examination showed decreased breath sounds and dullness to percussion on the posterior aspect of the left chest.
Chest x-ray showed a left-sided, alveolar consolidation with air bronchograms consistent with pneumonia and a large left-sided pleural fluid collection concerning for a parapneumonic effusion. The pertinent laboratory tests were as follows: 1) white blood cell count of $10.3 \times 10^3/\mu L$ (4.0 – $10.5 \times 10^3/\mu L$); 2) lactate dehydrogenase (LDH) 363 U/L (94 – 200 U/L); 3) ferritin 343.2 ng/mL (22.9 – 336.2 ng/mL); 4) fibrinogen 798 mg/dL (200 – 393 mg/dL); 5) D-dimer 1,144 ng/mL ($\leq$ 500 ng/mL); and 6) procalcitonin 0.16 ng/mL (0.00 – 0.50 ng/mL). Chest Computed Tomography (CT) demonstrated a large left pleural effusion, and mediastinal lymphadenopathy.

A therapeutic thoracentesis with ultrasound was performed with complete drainage and removal of 1200cc of serosanguinous fluid. Pleural fluid culture and cytology were negative. The fluid met criteria for exudative pleural fluid and the total nucleated cell count was 993 (/mm$^3$) with 89% lymphocytes. A RT-PCR (Xpert® Xpress SARS-CoV-2, Cepheid, Sunnyvale, CA) was performed on the pleural fluid and was positive for SARS-CoV-2. Sputum and blood cultures were negative. He was treated supportively for Acute Respiratory Distress Syndrome with high flow nasal cannula, dexamethasone and hydroxychloroquine. There was no recurrence of the pleural effusion during his hospital course.

Discussion:

To our knowledge, this is a novel description of COVID-19 related pleural effusion describing the pleural fluid analysis of virus detection confirmed by RT-PCR and behaving as a complicated parapneumonic effusion. There is a paucity of data describing the pleural complications in relation to the viral infections in general. A prior study by Sahn and colleagues$^1$, reported an 8% incidence of pleural effusions in
patients with acute viral infection, while using lateral decubitus chest x-rays incidence is reported to be 18%. However, the true incidence of viral related pleural effusion may be even higher if pleural ultrasonography is used as a screening tool for pleural fluid presence. In a review by Nestor and colleagues\textsuperscript{2}, the pleural fluid analysis in viral related pleural effusion is variable and typically exudative. Pleural fluid appearance can range from serous, serosanguineous, to hemorrhagic.

With respect to the SARS-CoV-2 pneumonia, review of literature reveals the prevalence of pleural effusion can be as high as 14\% in patients.\textsuperscript{3-5} In one of the largest trials, Li and colleagues described intrathoracic lymph node enlargement in 67\% of cases and pleural effusion in 14\% of cases.\textsuperscript{3} Patients with more severe cases have a higher prevalence of both lymphadenopathy and pleural effusions.\textsuperscript{3-5}

Pleural fluid analysis in this case showed an exudative effusion with high lymphocyte count. Even though pleural fluid analysis is rarely diagnostic in isolation, the differential diagnosis of a lymphocytic, exudative pleural effusion is broad and would include viral related pleural effusions. In our case, SARS-CoV-2 RNA was detected using RT-PCR, which is consistent with the diagnosis of a complicated parapneumonic effusion from SARS-CoV-2 related pneumonia.
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Figure Legend:
Figure 1: An axial image of a contrasted Chest Computed Tomography at the level of the bifurcation of the main pulmonary artery and at the level of the hilum. The chest CT demonstrates a large, left-sided, non-complex pleural effusion. There are pathologically enlarged lymph nodes, defined by >1cm in short axis, located in the sub-carinal and hilar regions.
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