Abstract

The 2009 pandemic H1N1 is a new virus. There is very little information related to the progression of the Influenza H1N1 2009 virus in immunosuppressive patients with a renal transplantation history. We presented the therapeutical management and clinical course of a patient who referred to our clinic because of pneumonia and respiratory failure due to Influenza H1N1 2009 virus and who received immunosuppressive therapy after renal transplantation.

Keywords: Renal transplantation; Influenza A H1N1 2009; Pneumonia

Introduction

The 2009 pandemic H1N1 is a new virus. Genetically, it is a mongrel, coming from three recognized sources (pigs, birds, and humans) which were combined in pigs [1]. In late March and early April 2009, an outbreak of H1N1 influenza A virus infection was detected in Mexico, with subsequent cases observed in many other countries [2]. On June 11, 2009, the World Health Organization raised its pandemic alert level to the highest level, phase 6, indicating widespread community transmission on at least two continents [3].

The risk of developing complications in patients with a medical state of an underlying disorder such as asthma, chronic obstructive pulmonary disease (COPD), immunosuppression, diabetes, obesity or chronic cardiac disease and children below 5 and patients over 65 years of age and pregnant women is higher and may require immediate hospitalization [4]. Patients who require intensive care generally may have a lung graph compatible with a hypoxia state and acute respiratory distress syndrome [5]. There is very little information related to the progression of the Influenza H1N1 2009 virus in immunosuppressive patients with a renal transplantation history.

We have presented the therapeutical management and clinical course of a patient who referred to our clinic because of pneumonia and respiratory failure due to Influenza H1N1 2009 virus and who received immunosuppressive therapy almost 7 years ago after renal transplantation.

Case Report

A 35-year-old male patient who underwent renal transplantation in 2002 due to chronic renal failure with an unknown etiology referred to our emergency service because of symptoms as high temperature, shortness of breath, dry cough, sore throat and headache and diffuse muscle pain. The complaints of this patient began suddenly nearly 24 hours before his referral to our clinic. Patient was receiving cyclosporine-A 2 x 100 mg, azathioprine 2 x 50 mg/day, prednisolone 1 x 10 mg/day, doxazosin 2 x 4 mg/day, diltiazem 1 x 60 mg/day and amlodipine 1 x 10 mg/day at the time of his referral. Patient was highly confused during physical examination and showed the following findings: number of respiration: 32/min, blood pressure: 170/110 mmHg, temperature: 38.8 °C, pulse rate: 116/min and oxygen saturation: 83% at ambient air. Dermal turgor was reduced, the oropharynx was slightly hyperemic and rales were present at bilateral lung basals. Heart beat was rhythmic and tachycardia was present.

Blood analysis revealed the following values: hemoglobin: 9.2 gr/dl, WBC: 9300/μl, Plt: 228000/μl, blood sugar:
98 mg/dl, urea: 67 mg/dl, creatinin: 1.72 mg/dl, AST: 18 U/L, ALT: 21 U/L, LDH: 392 U/L, CPK: 72 U/L, Na: 131 mmol/L, K: 3.4 mmol/L, Ca: 8.2 mg/dl, P: 4.3 mg/dl and CRP: 195 mg/L. Urinalysis was normal. Chest X-ray showed diffuse ground-glass density consolidation at the lower zone of the right lung, and a minimal ground-glass density at the lower zone of the left lung (Fig. 1). The patient was immediately transferred to the intensive care unit with pneumonia possibly due to H1N1 virus, confused consciousness and low oxygen saturation. Patient was somnolent and temperature was 38.8 °C. Oseltamivir 2 x 75 mg, azithromycin 500 mg/day and ceftriaxone 2g/day were initiated. We reduced the dosage of an immunosuppressive agent, azathioprine to 50 mg and increased the dose of prednisolon to 15 mg. Oxygen saturation in ambient air was 85% with 5 L/min nasal oxygen. We decided that there was no need for mechanical ventilation. Patient was followed up with nasal oxygen. However, acute renal failure developed during the stay at the intensive care unit and creatinine level increased to 2.7 mg/dl. Urine output, hydration status and bio-chemical values of the patient were carefully monitored and necessary liquid and electrolyte therapies were provided. Cyclosporine level was at the therapeutical range (Co: 102 mg/L). At day 5 in the intensive care unit, patient’s general condition was moderately good, consciousness state was open, and the oxygen saturation in ambient air was 88%. After administration of 3 L/min nasal oxygen, oxygen saturation increased to 93%. The following values were obtained: blood pressure: 140/90 mmHg, pulse: 95/min and temperature: 37.2 °C. At day 7, patient’s condition state improved and oxygen saturation at ambient air was measured as 92% and chest X-ray lung graphs and renal functions turned to normal and the patient was transferred back to a normal room (Fig. 2).

Definitive diagnosis of H1N1 was confirmed after discharged from hospital by nasal swab result which was done by Refik Saydam National Public Health Agency, Virology reference and research laboratory, national influenza center. Influenza A (H1N1)v PCR was performed with Qiagen artus

**Discussion**

Influenza A H1N1 2009 virus can spread out by direct contact as in seasonal flu and by the inhalation of infected airborne particles or by the means of droplet infection [6]. Typical clinical symptoms are fever, tremor, headache, coughing, a sore throat, myalgia and weakness. During the 2009 pandemic, it was reported that some patients experienced a rapidly progressive pneumonia, respiratory failure, acute respiratory distress syndrome and pulmonary emboli [7].

The patient presented in this article stated that his symptoms began 24 hours ago. Patient referred to our clinic with a pneumonia table. The clinical course was rapid and within 24 hours the patient was transferred to the intensive care unit. After onset of an early therapy regime, patient was discharged from the hospital within ten days due to a significant improvement in his general condition.

There are a small number of case presentations in the literature related to immunosuppressive patients with a developed H1N1 2009 virus infection after organ transplantation. Watcharananan et al [8] have reported the course of a H1N1 infection in two immunosuppressive patients. The first patient who underwent HD because of chronic rejection after renal transplantation and under low-dose maintenance steroid treatment referred to a clinic due to coughing and shortness of breath. An empiric therapy was initiated to this patient even though rapid antigen test was negative. Unfortunately, a few days later the patient’s general condition worsened and oral oseltamivir therapy was initiated due to the sudden requirement for non-invasive ventilation. Clinic healing was
observed and the patient was discharged from the hospital at Day 17. The other patient was receiving immunosuppressive agents due to renal transplantation and dyspnea developed at Day 2 after hospitalization to undergo a cataract operation. Considering the clinical status of the patient, oseltamivir 2 x 75 mg/day was immediately initiated even though the results of the rapid antigen test were negative. Eventually the dosage of oseltamivir was instantly increased. The patient was discharged 2 weeks later. Troppmann et al [9] reported a patient who referred to a hospital due to dyspnea and coughing complaints at Day 11 after renal transplantation. This patient failed to respond to treatment and died.

The rapid antigen test, direct fluorescent antibody test, PCR and culture test can be used in the diagnosis of influenza [10]. The 2009 H1N1 virus is sensitive to neuraminidase inhibitors such as oseltamivir [11]. According to the studies conducted, at an early stage antiviral treatment after infection may reduce the intensity and period of the disease and possibly provide early inhibition of the influenza virus replication as well [12]. Death reports related to the pandemic virus demonstrated the fact that delayed hospitalization and antiviral therapy were the main factors that may lead to a fatal result [13]. However, empiric therapy must be initiated in patients who are highly suspected to be infected by the virus based on their clinic table, even though results of tests are negative [14]. We preferred treating our patient with oseltamivir while PCR results were pending.

As a conclusion, symptoms of Influenza A H1N1 2009 virus in immunosuppressive patients may resemble healthy adults, however the clinical progression in immunosuppressive patients can be rapid and severe. In our case and as it is in other cases in the literature, 24 hours intensive care may be required. The success of early and high dose antiviral therapy is high and therefore starting therapy can be a life-saving factor even though a clinical doubt is present and a solid confirmation is not yet provided.

References