

COMMENTARY

# Rhinovirus-Infected Patients in the COVID-19 Pandemic Period

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Abstract: Human rhinovirus (HRV) is known as one of the most important respiratory pathogens, and the clinical characteristics of HRV infection might be similar to those of coronavirus disease 2019 (COVID-19). We identified 11 HRV-infected patients by polymerase chain reactions of the HRV genes among 151 outpatients with fever. All nine adult patients had underlying diseases and finally improved with the appropriate treatment in this COVID-19 pandemic period. Differential diagnosis between HRV and severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) infection will be needed to save lives and medical resources.

Keywords: SARS-CoV-2, human rhinovirus, PCR, antimicrobial stewardship

# **Commentary**

Since February 2020, the coronavirus disease 2019 (COVID-19) pandemic has been a huge issue in the world including Japan, and many clinics and hospitals have started polymerase chain reaction (PCR)-based surveillance of febrile patients and/ or patients with a history of close contact with COVID-19 persons and who visited a pandemic metropolitan area.<sup>1,2</sup> In this pandemic era, the trends of the other respiratory viruses, including influenza (flu), respiratory syncytial virus (RSV), human metapneumovirus (hMPV), and human rhinovirus (HRV) may be important factors affecting the prevalence of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2), because virus-virus interference is well known.<sup>3</sup>

In our hospital, we identified 11 HRV-infected patients among 151 febrile outpatients from October 10 to December 25, 2020 (Table 1). These patients were initially suspected to have COVID-19, but SARS-CoV-2 infection was not detected by PCR-based surveillance (cobas SARS-CoV-2 test, Roche, Basel, Switzerland).<sup>4</sup> We confirmed HRV infection using virus-specific primers (LightMix Modular Rhinovirus, Roche),<sup>5</sup> and the patients showed the characteristic clinical features: all nine "adult" patients had underlying diseases, although the other three "pediatric" patients (Patients 5, 8, and 9) had been healthy. A variety of diseases were diagnosed, and bacteria, such as Escherichia coli, Streptococcus pneumoniae, and Staphylococcus aureus were isolated from some patients (Patients 2, 4, and 5), but they finally improved with the appropriate treatments, including antibiotics, such as meropenem, ampicillin/sulbactam, and cefazolin. Only one parainfluenza virus-infected patient was found (data not shown), but no other viruses, including Flu, RSV, hMPV, and SARS-CoV-2, were detected among the 151 febrile outpatients.

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Table I Clinical Characteristics of Human Rhinovirus Infected Patients

Age	Male/Female	Male/Female Detected Virus	Onset Location	Underlying Diseases	Final Diagnosis	اemp ر	Chest Shadows	Treatment	Improvement	Others/Sample
83	ட	Rhinovirus	Nursing home	Parkinson diseases	Common cold	37.6	Slightly	Acetaminophen	Yes	None
89	Σ	Rhinovirus	Community	GB stone	Cholecystitis	38	Slightly	MEPM	Yes	E. coli/Blood
46	ш	Rhinovirus	Community	DM, Collagen	Nephritis	37	None	None	Yes	None
88	Σ	Rhinovirus	Nursing home	CKD, HD	Pneumonia	38	Strong	ABPC/SBT	Yes	S. pneumoniae Ag/Urine
2	Σ	Rhinovirus	Community/Home	None	Cellulitis	37.2	None	CEZ	Yes	S. aureus/Skin
37	ш	Rhinovirus	Hospital	Collagen	Cellulitis	39	None	AMPC	Yes	None
30	Σ	Rhinovirus	Community	SAS, Atopy	Common cold	39	None	Acetaminophen	Yes	None
15	ш	Rhinovirus	Community	None	Common cold	38	None	Mucodain	Yes	None
12	ш	Rhinovirus	Community	None	Otitis media	38	None	Acetaminophen	Yes	None
53	ш	Rhinovirus	Community	SLE	Bronchiolitis	38	Slightly	LVFX	Yes	None
87	Σ	Rhinovirus	Community	Prostate cancer	Common cold	37.2	Slightly	AMPC	Yes	None

HRVs are known to be the responsible pathogens for more than half of the cold-like illnesses, and recently they have been found not only as upper respiratory tract pathogens, but also as lower bronchial tract pathogens, in particular, in patients with asthma, infants, elderly patients, and immunocompromised hosts.<sup>6</sup> The symptoms, signs, and pathogenesis of HRV infection may be severe and similar to COVID-19 in patients with underlying diseases. Therefore, accurate differential diagnosis is critical to save the patients and valuable medical resources, including antimicrobial agents, based on the antiviral stewardship philosophy. In addition, the development of anti-HRV agents may also be needed to improve these patients' conditions quickly in the COVID-19 pandemic era.

### **Ethics**

The analysis was approved by the Committee for Clinical Scientific Research of Tohoku Medical and Pharmaceutical University Hospital on May 21, 2017 (No. ID 2017-2-010), and the patients provided written, informed consent for use of their specimens, although the samples were collected as part of routine laboratory analyses.

## **Disclosure**

The authors report no conflicts of interest in this work.

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ulbactam; CEZ, cefazolin; AMPC, amoxicillin; LVFX, levofloxacin

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