DOI: 10.1002/ams2.854

### LETTER TO THE EDITOR

# Beware of accidental ingestion of *Colchicum autumnale* mistaken for *Allium victorialis*

Dear Editor,

According to the statistical data on food poisoning from the Ministry of Health, Labor, and Welfare, two cases of colchicine poisoning occurred in 2021, resulting in one death. Additionally, three more such cases occurred in 2022, resulting in two deaths, indicating that lethal cases of this type of poisoning still occur.<sup>1,2</sup> We encountered a fatal case of colchicine poisoning because of the accidental ingestion of *Colchicum autumnale*, and we describe it here.

An 86-year-old man with a history of diabetes mellitus, chronic heart failure, atrial fibrillation, severe tricuspid regurgitation, postoperative gastric cancer status, and irondeficiency anemia had accidentally ingested *Colchicum autumnale* (including the stem at the base and 2 to 3 leaves, but no bulbs), which had grown in his garden in the form of tempura. The patient developed diarrhea the day after ingestion. However, he refused hospitalization and was observed at home. Three days after the ingestion, the patient experienced difficulty moving and was taken by ambulance to a nearby hospital.

On admission, he had developed multiple organ failure, including circulatory shock, acute liver failure, acute kidney injury (AKI), and coagulation disorders. The patient was transferred to our hospital for critical care. He presented with a consciousness level corresponding to a score of 12 on the Glasgow Coma Scale (E4V2M6) and was administered noradrenaline. On physical examination, his blood pressure was 137/64 mm Hg, heart rate was 98 beats per minute, respiratory rate was 28 breaths per minute, and body temperature was 37.1°C. The results of his blood tests were as follows: white blood cell count of 9600/µL, hemoglobin of 13.7 g/dL, and platelet count of  $11.8 \times 10^4$ / µL. The results of the coagulation tests were prothrombin time-international normalized ratio (PT-INR), 3.56; D-dimer, 177.0 µg/mL; fibrin degradation product (FDP), 433.0 µg/mL; and an acute disseminated intravascular coagulation (DIC) score of 6 points. Biochemical tests revealed: aspartate aminotransferase (AST), 635 U/L; alanine aminotransferase (ALT), 250 U/L; lactate dehydrogenase (LD IFCC), 2294 U/L; alkaline phosphatase (ALP IFCC), 404 U/L; creatine kinase (CK), 1216 U/L; blood urea nitrogen (BUN), 50.8 mg/dL; creatinine (CRE), 3.30 mg/ dL; C-reactive protein (CRP), 10.48 mg/dL; and blood

sugar, 66 mg/dL. The results of an arterial blood gas analysis, which was collected on use of an oxygen mask at 10 L/ min, indicated a pH of 7.24,  $PaO_2$  of 105 mm Hg,  $PaCO_2$ of 58 mm Hg,  $HCO_3^-$  of 24.9 mmol/L, and a lactate level of 77 mg/dL. The family of the patient brought in samples of the *Colchicum autumnale* that had been mistakenly ingested (Figure 1). After confirming that it was not *Allium victorialis*, activated charcoal was administered to the patient via a nasogastric tube, after the placement of a cuffed endotracheal tube. Blood purification therapy was initiated for the management of acute liver failure and AKI. Despite intensive care unit treatment, which included artificial ventilation support and acute blood purification therapy, his condition rapidly deteriorated, and he died on the fourth day after the accidental ingestion.

Colchicine strongly inhibits cell division and intracellular metabolism and can damage cells and organs with fast metabolic turnover, such as small intestinal mucosal cells and bone marrow cells. The half-life of colchicine is 10 to 30 min and the time to reach peak blood concentration is 30 min to 2h. However, the inhibitory effect on cell division reaches its peak ~10h after ingestion, with a latency period of 2 to 12h until toxic symptoms appear. Although no specific effective treatment exists, symptomatic therapy is necessary to overcome the acute phase of multi-organ failure, which lasts for ~1 week. A level of 0.3 to 2.5 ng/mL of colchicine in the blood is considered therapeutic, that of 5.0 ng/mL or higher is considered toxic, whereas that of 9.8 ng/mL or higher is considered lethal.<sup>3</sup>

In our case, the colchicine blood concentration remained elevated, with levels of 5.0 and 4.0 ng/mL on the third and fourth day after ingestion, respectively. The colchicine concentration in *Colchicum autumnale* samples obtained in this study was higher than those previously reported by the Akita Prefecture, with concentrations of 2028.7, 1873.7, and 2004.6  $\mu$ g/g in the leaves, stems, and roots, respectively. Notably, colchicine was not detected in *Allium victorialis* samples.<sup>4</sup> The minimum lethal dose of colchicine is 4.3 mg in a person weighing 50 kg (86  $\mu$ g/kg). Additionally, a colchicine concentration of 2028.7  $\mu$ g/g in the leaves of the plant translates to a dose of 2.1 g, which is sufficient to reach the minimum lethal dose in humans. Therefore, the *Colchicum autumnale* has high toxicity, resulting in

This is an open access article under the terms of the Creative Commons Attribution-NonCommercial License, which permits use, distribution and reproduction in any medium, provided the original work is properly cited and is not used for commercial purposes.

© 2023 The Authors. Acute Medicine & Surgery published by John Wiley & Sons Australia, Ltd on behalf of Japanese Association for Acute Medicine.

# ACUTE MEDICINE WILEY-& SURGERY





Allium victorialis

# Colchicum autumnale

**FIG. 1** *Colchicum autumnale* collected at the patient's home and commercially available *Allium victorialis*. Colchicine concentration in the leaves, stems, and roots of *Colchicum autumnale* is of 2028.7, 1873.7, and 2004.6 µg/g, respectively. Contrarily, colchicine was not detected in *Allium victorialis* samples.

multiple organ failures and mortality.<sup>5</sup> Colchicine poisoning manifests with non-specific symptoms and can only be diagnosed with the aid of a detailed medical history and assessment of the severity of symptoms. Healthcare professionals should be aware of *Colchicum autumnale* toxicity and public education on the prevention of accidental ingestion is warranted.

## ACKNOWLEDGMENTS

We express our deep gratitude to the members of the Education and Research Center of Legal Medicine, Chiba University, for their assistance in the measurement of colchicine blood concentration and to the members of the Akita Prefectural Research Center for Public Health and Environment for their contribution to the identification of *Colchicum autumnale*. We thank Editage (www.edita ge.com) for English language editing.

## CONFLICT OF INTEREST STATEMENT

Authors declare no conflict of interests for this article.

#### ETHICS STATEMENT

Approval of the Research Protocol: Not applicable.

Informed Consent: Informed consent for publication was obtained from the patient's family.

Registry and Registration No. of the Study/Trial: Not applicable.

Animal Studies: Not applicable.

Tasuku Nara Hajime Nakae Yasuhito Irie Koumei Kameyama Manabu Okuyama

Department of Emergency and Critical Care Medicine, Akita University Graduate School of Medicine, Akita, Japan

#### Correspondence

Hajime Nakae, Department of Emergency and Critical Care Medicine, Akita University Graduate School of Medicine, 1-1-1 Hondo, Akita 010-8543, Japan.

Email: nakaeh@doc.med.akita-u.ac.jp

## ORCID

Hajime Nakae b https://orcid.org/0000-0003-3733-3530 Yasuhito Irie b https://orcid.org/0000-0002-1628-9262

#### REFERENCES

- Ministry of Health. Labour and Welfare [Internet]. Food Poisoning. [cited 2023, March 10]. Available from: https://www.mhlw.go.jp/stf/ seisakunitsuite/bunya/kenkou\_iryou/shokuhin/syokuchu/04.html
- Yokoyama T, Otsuki I, Kimijima T, Kawaguchi R, Takakuwa K, Nakabayashi K. A case of cardiomyopathy in the course of colchicine poisoning due to accidental ingestion of *Colchicum autumnale*. J Jpn Soc Intensive Care Med. 2022;29:43–4.

- Schulz M, Schmoldt A, Andresen-Streichert H, Iwersen-Bergmann S. Revisited: therapeutic and toxic blood concentrations of more than 1100 drugs and other xenobiotics. Crit Care. 2020;24:195. https://doi. org/10.1186/s13054-020-02915-5
- 4. Fujii M, Matsubuchi A, Furui M, Nakamura J, Wakasa N. Analysis of colchicine and other compounds in prepared dishes of a toxic plant,

*"Colchicum autumnale"*, using LC-MS/MS. Ann Rep Akita Prefectural Res Center Public Health Environ. 2021;17:51–2.

 Giorgetti A, Nalesso A, Cecchetto G, Pizzi M, Bellan A, Viel G, et al. Two fatal intoxications by colchicine taken for saffron. Clinical, medico-legal and forensic toxicological implications. Leg Med. 2019;38:64–8.