



Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.



Figure 1. Several photographs were reviewed for each infant after various adverse food reactions throughout OIT. This figure reveals case 3 infant at 9 months of age when she developed a diffuse urticarial rash and bilateral eyelid swelling associated with rhinorrhea without additional symptoms within minutes after ingestion of several bites of cashew butter with no contact exposure. She was given diphenhydramine, and symptoms resolved within hours. OIT, oral immunotherapy.

unresponsiveness. This case series is limited by the small sample size and will require validation in a larger cohort. The medical background of these parents substantially limits the generalizability of this case series and the lack of comprehensive testing limits our understanding of the degree of sensitization to food allergens and may call into question the underlying food allergy diagnosis. In addition, the long-term outcomes of food desensitization remain unknown.

Given the global burden of IgE-mediated food allergies, strategies to mitigate the overall morbidity and mortality associated with this disease are critical. This limited case series should not be used to change clinical practice. Rather, we hope that sharing these stories will encourage clinics with more robust data on infant OIT to come forward, to better define the risks, and to allow informed decision-making for both physicians and parents. This case series highlights a need for more research exploring the safety and efficacy of infant OIT.

Hannah Wangberg, MD*

Laura Kuhlman, MD†

Kevin Cook, MD*

* Division of Allergy, Asthma, and Immunology
Scripps Health
San Diego, California

† Department of Emergency Medicine
Indiana University Health, Inc
Muncie, Indiana

Wangberg.Hannah@scrippshealth.org

References

1. Sicherer SH, Sampson HA. Food allergy: a review and update on epidemiology, pathogenesis, diagnosis, prevention, and management. *J Allergy Clin Immunol.* 2018;141(1):41–58.
2. Gupta RS, Springston EE, Warrier MR, et al. The prevalence, severity, and distribution of childhood food allergy in the United States. *Pediatrics.* 2011;128(1):e9–e17.
3. Osborne NJ, Koplin JJ, Martin PE, et al. Prevalence of challenge-proven IgE-mediated food allergy using population-based sampling and predetermined challenge criteria in infants. *J Allergy Clin Immunol.* 2011;127(3):668–676. :e1–2.
4. Roehr CC, Edenharter G, Reimann S, et al. Food allergy and non-allergic food hypersensitivity in children and adolescents. *Clin Exp Allergy.* 2004;34(10):1534–1541.
5. Du Toit G, Roberts G, Sayre PH, et al. Randomized trial of peanut consumption in infants at risk for peanut allergy. *N Engl J Med.* 2015;372(9):803–813.
6. Du Toit G, Sampson HA, Plaut M, Burks AW, Akdis CA, Lack G. Food allergy: update on prevention and tolerance. *J Allergy Clin Immunol.* 2018;141(1):30–40.
7. Perkin MR, Logan K, Bahnson HT, et al. Efficacy of the Enquiring About Tolerance (EAT) study among infants at high risk of developing food allergy. *J Allergy Clin Immunol.* 2019;144(6):1606–1614. e2.
8. US Food and Drug Administration. FDA approves first drug for treatment of peanut allergy for children. Available at: <https://www.fda.gov/news-events/press-announcements/fda-approves-first-drug-treatment-peanut-allergy-children>. Accessed December 30, 2020.
9. O'B Hourihane J, Beyer K, Abbas A, et al. Efficacy and safety of oral immunotherapy with AR101 in European children with a peanut allergy (ARTEMIS): a multicentre, double-blind, randomised, placebo-controlled phase 3 trial. *Lancet Child Adolesc Health.* 2020;4(10):728–739.
10. Vickery BP, Berglund JP, Burk CM, et al. Early oral immunotherapy in peanut-allergic preschool children is safe and highly effective. *J Allergy Clin Immunol.* 2017;139(1):173–181. e8.

Skin prick test practice in allergy clinics during coronavirus disease 2019 pandemic



Skin prick test (SPT) is a vital tool to confirm sensitization in allergic diseases, making it a daily routine for allergists. Although SPT is not considered an aerosol-generating procedure, the close contact between the patient and the performer may pose a risk for transmitting the disease.¹ For Turkish health care facilities, the first phase of the coronavirus disease 2019 (COVID-19) pandemic started with identifying the first case and continued by a quarantine period (QP), declared at the end of March 2020. At the beginning of June 2020, the

Turkish government initiated the second period and named it the “normalization period” (NP).² We aimed to compare the situation and behavior of allergists between the 2 phases of the COVID-19 pandemic.

This study is an online survey-based, cross-sectional study conducted between June 20, 2020, and June 28, 2020. Participants were recruited through e-mails and WhatsApp. To our knowledge, this survey presents the first data of allergists' approaches to performing SPTs during the COVID-19 pandemic. The health care system in Turkey is mainly socialized. Turkey is administratively divided into 81 provinces, but only 33 have at least 1 allergist working. The survey

Disclosures: The authors have no conflicts of interest to report.

Funding: The authors have no funding sources to report.

was sent to nearly 300 allergists, and 128 responded by filling the questionnaire; so, we believe that the results of this study are sufficient to represent our country. The participants were from 28 different provinces in Turkey. A total of 55 of the respondents (43%) were pediatric allergists. In the QP, nearly half of the participants had to shut down the SPT laboratories and one-third of them had to work in pandemic outpatient clinics. The ratio of participants who continued to perform SPTs in the QP was 54%. Of all the participants, 16% stated that their prick test laboratories were moved to a different place in their hospital. In the QP, 60% of the participants stated that they were very concerned on the risk of transmission during SPT. Despite wearing masks, 60% of the participants were anxious on being face-to-face with a patient for 15 minutes. Hospital and personal phones were the most preferred methods of telecommunication. There were 43% of the participants who preferred to perform all SPTs by appointment, 36% suggested a partial appointment system, whereas 21% did not consider it necessary to have an appointment system. In addition, 90 (71%) participants considered that the test intervals should be at least 30 minutes. When asked on the diagnostic value of the SPT for the treatment of allergy during the pandemic, 29% answered SPT to be very important, whereas 51% of the participants considered it of moderate importance. Of the participants, 76 (60%) stated that they would prefer to order specific immunoglobulin E tests more than they usually do. Furthermore, 87 participants said that they would prioritize testing with venom, 71 with food allergens, and 49 with latex (Fig 1). Priority given to food allergens was 65% among pediatric allergists and 34% among adult allergists. With the transition to the NP, the ratio of participants who had high anxiety on performing an SPT decreased from 60% to 30%.

The European COVID-19 outbreak position document recommends that SPT should be generally suspended or replaced by laboratory tests during the pandemic except for some individual cases.¹ In our questionnaire, most participants attributed importance to venom allergy and, especially, pediatric allergy specialists gave more priority to food allergens by stating that they would continue to perform skin prick tests with these allergens despite the pandemic. Owing to the urgent nature of these allergies, we consider this result an excellent and practical example of a careful risk-benefit assessment, which the European Academy of Allergy and Clinical Immunology and Allergic

Rhinitis and its Impact on Asthma has mentioned.¹ More than half of the participants preferred to perform an SPT at least 30 days after treatment completion of COVID-19 in a patient. In other words, participants did not want to test even if the official quarantine period was over.

One consequence of the pandemic is the increased exposure to disinfectants. The American Nurses' study revealed that poor asthma control was associated with exposure to formaldehyde, hypochlorite bleach, and hydrogen peroxide but not with exposure to alcohol-based disinfectants.³ Most participants chose chlorine-containing disinfectants for the floors and alcohol-based disinfectants for the surfaces of seats and desks. We consider this result to be according to the recommendations of the Centers for Disease Control and Prevention that advise patients to choose disinfectants that are less likely to cause an asthma attack, such as products with ethanol (ethyl alcohol).⁴

We found that only one-third of the participants had a written action plan. This low result is likely to be related to the fact that the survey was conducted at the beginning of the pandemic for our country. The World Health Organization shares daily information on the number of coronavirus cases for countries.⁵ Following this information may help adjust allergy clinics by predicting the pandemic course in the upcoming time. Basic reproductive rate (R_0) is the average number of people infected by 1 person in a susceptible population.⁶ For Turkey, R_0 was below 1 in the NP, and we revealed in our study that physicians felt less anxious when performing an SPT during this period. An implication of our result to clinical practice may be the possibility of approaching SPTs according to the R_0 ratio. In this sense, an R_0 ratio below 1 may refer to continuing normal operations. In contrast, an R_0 ratio above 1 may refer to postponing SPTs similar to the stratified approach mentioned by Shaker et al⁷ in a special article on pandemic contingency planning for allergy clinics.

Our suggestion, because it would not be practical to suspend all SPTs for long periods, is that the decision on prioritization of testing should be made carefully. For instance, SPT with venom and food extracts may be done more urgently than others. Proper measures, such as universal masking and effective triage, to create a safer environment are needed. Performing SPTs by appointment would be reasonable. The primary limitation of this study is the lack of effective, sustainable, or verified data backing up our

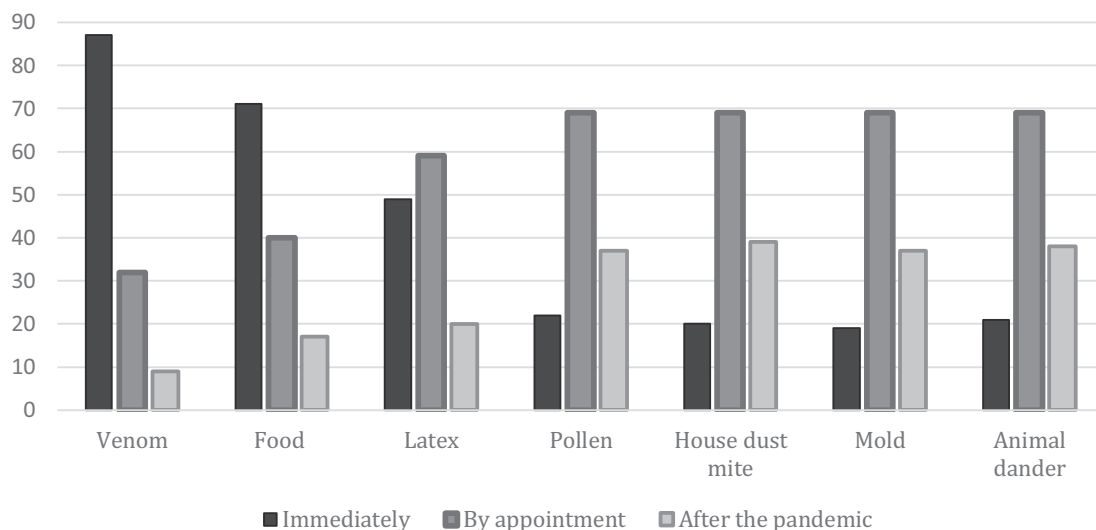


Figure 1. The answers of the respondents to the "Please select your timing preference for performing a skin prick test regarding allergens during the pandemic." The top priority was attributed to the venom allergen ($n = 87$), followed by food allergens ($n = 71$) and latex ($n = 49$).

findings. Therefore, we suggest that further studies are needed to validate our recommendations.

resat.kendirlinan@saglik.gov.tr

Resat Kendirlinan, MD^{*,†}

Pamir Cerci, MD[‡]

Dilsad Mungan, MD[§]

^{*} Division of Immunology and Allergy

Department of Chest Diseases

Aydın Government Hospital

Aydın, Turkey

[†] Division of Immunology and Allergy

Department of Chest Diseases

Izmir Katip Celebi University Atatürk Education and

Research Hospital

Izmir, Turkey

[‡] Division of Immunology and Allergy

Department of Internal Medicine

Eskisehir City Hospital

Eskisehir, Turkey

[§] Division of Immunology and Allergy

Department of Chest Diseases

School of Medicine

Ankara University

Ankara, Turkey

References

1. Pfaar O, Klimek L, Jutel M, et al. COVID-19 pandemic: practical considerations on the organization of an allergy clinic—an EAACI/ARIA position paper. *Allergy*. 2021;76(3):648–676.
2. Guide on working in healthcare institutions during the normalization period in COVID-19 pandemic. Study by Scientific Advisory Board, Republic of Turkey Ministry of Health. 2020. Available at: <https://covidlawlab.org/wp-content/uploads/2020/07/Guide-to-working-in-healthcare-institutions-during-normalization-.pdf>. Accessed June 9, 2020.
3. Dumas O, Wiley AS, Quinot C, et al. Occupational exposure to disinfectants and asthma control in US nurses. *Eur Respir J*. 2017;50(4):1700237.
4. Centers for Disease Control and Prevention. People with moderate to severe asthma. Available at: <https://www.cdc.gov/coronavirus/2019-ncov/need-extra-precautions/asthma.html>. Accessed November 28, 2020.
5. World Health Organization. Infection prevention and control guidance for long-term care facilities in the context of COVID-19: interim guidance, 21 March 2020. Available at: https://apps.who.int/iris/bitstream/handle/10665/331508/WHO-2019-nCoV-IPC_long_term_care-2020.1-eng.pdf. Accessed November 29, 2020.
6. Li Y, Campbell H, Kulkarni D, et al. The temporal association of introducing and lifting non-pharmaceutical interventions with the time-varying reproduction number (R) of SARS-CoV-2: a modelling study across 131 countries. *Lancet Infect Dis*. 2021;21(2):193–202.
7. Shaker MS, Oppenheimer J, Grayson M, et al. COVID-19: pandemic contingency planning for the allergy and immunology clinic. *J Allergy Clin Immunol Pract*. 2020;8(5):1477–1488.e5.

Food protein–induced enterocolitis syndrome in monoamniotic twins



Food protein–induced enterocolitis syndrome (FPIES) is considered a non–immunoglobulin E (IgE)-mediated food allergy that typically presents in infancy. It is characterized by gastrointestinal symptoms, mainly protracted and delayed vomiting.¹ Globally, the most often reported FPIES triggers are cow's milk (CM), soy, and grains (including rice and oats).^{1,2} Nevertheless, common food triggers differ between countries; these variations may be because of several factors, including geography, racial and ethnic differences, ages of patients, FPIES phenotype, and probably genetic and epigenetic factors.³ FPIES has been barely reported in siblings,⁴ it does not seem to be a strong familial association, limited data are available regarding twins with FPIES.^{5,6}

We report an unusual case of FPIES in monoamniotic monoamniotic female twins. They were born at 32 weeks and 1 day of gestational age by cesarean delivery; both girls had atopic dermatitis and transient hypogammaglobulinemia of infancy (markedly for IgA and relative of IgM) that resolved after 6 months, with a quantitative study of lymphocyte population within normal limits.

They received exclusive breastfeeding until 8 months (6 months of corrected age) when they started the weaning process with gluten-free cereals and fruits, with good tolerance. At 9 months old, after eating a multigrain porridge (wheat, corn, rice, oats, barley, and rye), both twins developed, 2 hours after the intake, protracted vomiting (6 vomits) accompanied with marked paleness and lethargy; the symptoms resolved within 4 hours and they did not require emergency assistance. Gluten was removed from their diet, and they remained asymptomatic. They continued eating gluten-free cereals. After 1 month, their parents decided to reintroduce gluten cereals into the diet of their children. On the second time the twins ate the same

porridge, they developed repeated vomiting, paleness, and lethargy. Symptoms started 1 hour and a half after the ingestion and resolved within a couple of hours. An allergology study was carried out. Skin prick test (SPT) was performed with commercial extracts to wheat, barley, rye, and oats. Serum-specific IgE (sIgE) was measured with the ImmunoCAP system (Thermo Fisher Scientific, Uppsala, Sweden) with negative results. Serum detection of tissue anti-transglutaminase IgA and IgG and deamidated gliadin peptide antibody IgG was negative. Gluten-FPIES was suspected by a compatible clinical history: at least 2 reproducible episodes of gastrointestinal symptoms triggered by a specific food within 1 to 12 hours, absence of symptoms suggesting an IgE-mediated reaction, resolution of symptoms after elimination of the culprit food, and no other explanation for the symptoms identified. A gluten-free diet was indicated, no accidental exposures occurred, and other food groups (egg, CM, fish, fruits, meats) were introduced into the diet of the girls with good tolerance. They remained asymptomatic, and both twins had healthy general conditions and proper growth indicators. After 1 year, the allergology study was repeated. The SPT result remained negative, and an oral food challenge (OFC) with the porridge involved in the reaction was performed. The OFC was carried out in 2 nonconsecutive days, following our protocol⁷ (Fig 1). Both girls tolerated the normal serving size per age without developing symptoms.

FPIES is considered a rare disease; however, it seems not as rare as it is thought to be but remains underdiagnosed. The prevalence studies reported a cumulative incidence range between 0.015% and 0.7%.^{4,8} Unlike what happens with IgE-mediated allergy, FPIES does not seem to have a strong familial association. Mehr et al⁴ reported a prevalence of FPIES in 7% of the siblings included in their study, but limited data are available in the literature regarding twins with FPIES and the degree of concordance of the disease. Watanabe et al⁵ reported a case of CM-FPIES in monozygotic twin neonates. Feeding

Disclosures: The authors have no conflicts of interest to report.

Funding: The authors have no funding sources to report.