

UDC 616.248-002-053.2:613.16(477)

O. V. Mozyrska¹, Ya. V. Socolova²

The role of sensitization to aeroallergens and its determination in children with asthma in Ukraine

¹Bogomolets National Medical University, Kyiv, Ukraine²Kyiv City Children's Clinical Hospital No. 2, Ukraine

Modern Pediatrics. Ukraine. (2024). 6(142): 15-19. doi: 10.15574/SP.2024.6(142).1519

For citation: Mozyrska OV, Socolova YaV. (2024). The role of sensitization to aeroallergens and its determination in children with asthma in Ukraine. Modern Pediatrics. Ukraine. 6(142): 15-19. doi: 10.15574/SP.2024.6(142).1519.

Asthma is a very common respiratory disease in the world. In particular, asthma has a great impact on the health of children. In this study, the distribution of sensitization among children with asthma has been analyzed.

The aim of this study was to investigate the distribution of sensitization among children with asthma in the Ukrainian population.

Materials and methods. The study involved patients (n=98) with asthma aged from 12 to 18 years (median 6 (3; 10)). During the examination, children underwent skin prick testing for common aeroallergens. A positive skin prick test was defined as a blister ≥ 3 mm in diameter after 15 minutes.

Results. Among 98 patients, skin testing with aeroallergens showed a positive result in 76 (77.6%) people: 18 (18.4%) had positive skin testing for at least one allergen, and 58 (59.2%) had polysensitization. Among those sensitized children, 54 (55.1%) were sensitive to pollen allergens, 29 (29.6%) patients were sensitive to cats, 32 (32.6%) to dogs, 30 (30.6%) to *Dermatophagoides pteronyssinus*, 27 (27.6%) to *Dermatophagoides farinae*, and 10 (10.2%) to *Alternaria*.

Conclusion. The most important inhaled allergens for patients with bronchial asthma are pollens (trees, grasses, and weeds), house dust mites, molds, and animal dander. Identifying causative allergens allows for monitoring allergy triggers and determining which asthma patients will be candidates for allergen-specific immunotherapy (ASIT). Today, most children with asthma do not receive ASIT because of the lack of awareness at the primary care level, a lack of testing for aeroallergen sensitization, and financial inaccessibility of examination and treatment in Ukraine.

The study was carried out in accordance with the principles of the Declaration of Helsinki. The study protocol was approved by the Local Ethical Committee of these institutions. The informed consent of the children's parents was obtained for the research.

No conflict of interests was declared by the authors.

Keywords: asthma, aeroallergens, skin prick tests, children.

Роль сенсibilізації до аерoалергенів та її визначення в дітей із бронхіальною астмою в Україні

O. V. Мозирська¹, Я. В. Соколова²¹Національний медичний університет імені О.О. Богомольця, м. Київ, Україна²Київська міська клінічна лікарня №2, Україна

Бронхіальна астма є дуже поширеним респіраторним захворюванням у світі. Зокрема, бронхіальна астма має великий вплив на здоров'я дітей. У цій роботі було проаналізовано розподіл сенсibilізації серед дітей, хворих на бронхіальну астму.

Метою цього аналізу було дослідити розподіл сенсibilізації серед дітей, хворих на бронхіальну астму в Україні.

Матеріал і методи. У дослідженні взяли участь хворі (n=98) на бронхіальну астму віком від 12 до 18 років (медіана 6 (3; 10)). Під час обстеження дітям було проведено шкірне прік-тестування на поширені аерoалергени. Позитивний шкірний прік-тест визначався як пухир ≥ 3 мм у діаметрі через 15 хвилин.

Результати. Серед 98 пацієнтів шкірне тестування з аерoалергенами виявило позитивний результат у 76 (77,6%) осіб: у 18 (18,4%) — позитивна шкірна проба хоча б на один алерген, у 58 (59,2%) — полісенсibilізація. Серед сенсibilізованих 54 (55,1%) дитини були чутливими до пилоквіх алергенів, до кота — 29 (29,6%) хворих, собак — 32 (32,6%), *Dermatophagoides pteronyssinus* — 30 (30,6%), *Dermatophagoides farinae* — 27 (27,6%), альтернатію — 10 (10,2%).

Висновок. Найважливішими інгаляційними алергенами для хворих на бронхіальну астму є пилок (дерев, трав і бур'янів), кліщі домашнього пилу, цвіль і алергени тварин. Визначення причинних алергенів дає змогу контролювати тригери алергії та визначати, які пацієнти з астмою будуть кандидатами для алерген-специфічної імунотерапії (АСІТ). На сьогодні більшість дітей, хворих на бронхіальну астму, не отримують АСІТ через недостатню поінформованість на первинному рівні, відсутність алергологічного тестування, фінансову недоступність обстеження та лікування в Україні.

Дослідження виконано відповідно до принципів Гельсінської декларації. Протокол дослідження ухвалено Локальним етичним комітетом зазначеної в роботі установи. На проведення досліджень отримано інформовану згоду пацієнтів.

Автори заявляють про відсутність конфлікту інтересів.

Ключові слова: бронхіальна астма, аерoалергени, шкірні прік-тести, діти.

Introduction

Asthma is a very common respiratory disease in the world [16]. In particular, asthma has a great impact on the health of children [15,16]. The prevalence of asthma, like the prevalence of allergies, is increasing worldwide [14]. In Ukraine, the importance of asthma and allergies is also growing, as in a country with developed industrialization and the consequences of the Chernobyl disaster [2,4,17,18]. Military actions on the territory of the country can play a big role in the incidence, course, and control of the disease [19].

Due to the unavailability of examination for patients with asthma, the presence of allergies may remain undetected in many patients, and the impact of contact with the allergen may play a large role in the course of the disease [11,12].

Early diagnosis of asthma and establishment of sensitization and allergy can play a very important role in the treatment of the disease: allergen avoidance and allergen-specific immunotherapy [5,6]. Asthma guidelines or algorithms generally recommend the determination of specific IgE (sIgE) or skin prick testing for aeroallergen sensitization in subpopulations of patients with asthma [5,6].

A European primary care survey conducted by the European Academy of Allergy and Clinical Immunology (EAACI) and the International Primary Care Respiratory Group (IPCRG) found that 20.6% of practices had no access to allergy tests [1].

Aeroallergen sensitization testing increases the ability to predict the course of the disease, medication response, and the risk of future asthma exacerbations. Asthma Predictive Index (mAPI), which includes the role of allergic sensitization to at least 1 aeroallergen, is an important predictive model for childhood asthma. In the high-risk cohort, a positive mAPI significantly increased the likelihood of developing asthma in the future [4]. Allergy testing results can guide clinical precision medicine for chronic airway disease in individual patients, including allergen-specific immunotherapy (ASIT). In addition, allergy testing can help identify the presence of allergic rhinitis (AR), a common comorbidity, which, if properly treated, can improve asthma control [5,8].

The aim of this study was to investigate the distribution of sensitization among children with asthma of the Ukrainian population.

Materials and methods of the study

The study involved patients (n=98) with asthma aged from 12 to 18 years (median 6 (3; 10)), from the allergological department of the Kyiv City Children's Clinical Hospital No. 2.

The main group consisted of 82 boys and 16 girls, median age – 15 (11; 16).

The diagnosis of asthma was established in the group of examined children on the basis of clinical data and objective measurement of lung function [7]. All examined children underwent spirometry according to the ATS-ERS guidelines [9]: at least four acceptable maneuvers after the spirometers were checked for volume accuracy. The diagnosis of asthma was made if parents reported symptoms of asthma (difficulty breathing, chest tightness in the past 12 months, difficulty breathing or wheezing after exercise, wheezing in the previous 12 months, or a previous physician diagnosis of asthma and the child demonstrated significant airway reversibility (>12% increase in FEV₁).

The diagnosis of AR was determined in 80 (81.6%) patients on the basis of clinical history, positive physical examination, and the presence of one or more following nasal symptoms: rhinorrhea, nasal congestion, sneezing and itching for 3 months. Among them, 24 (30 %) had symptoms of year-round AR, and 56 (70 %) – had symptoms only in the pollen season. 33 (41.3 %) patients reported symptoms related to contact with animals.

During the examination, children underwent skin prick testing for aeroallergens (birch (*Betula verrucosa*), alder (*Alnus glutinosa*), hornbeam (*Capinus betulus*), hazel (*Corylus avellana*), timothy grass (*Phleum pratense*), ryegrass (*Lolium perenne*), orchard grass (*Dactylis glomerata*), ragweed (*Ambrosia Artemisiifolia*), mugwort (*Artemisia vulgaris*), sunflower (*Helianthus annuus*), common couch (*Elytrigia*) and *Alternaria* (*Alternaria alternate*), cat, dog, and dust mites (*Dermatophagoides farinae* and *Dermatophagoides pteronyssinus*). A positive skin prick test was defined as a blister ≥ 3 mm in diameter after 15 minutes.

Statistical processing of the obtained data was carried out using the statistical package IBM SPSS Statistics Base (version 22) and the software EZR version 1.32 (graphical interface of the R environment, version 2.13.0). The research database was systematized in the Microsoft

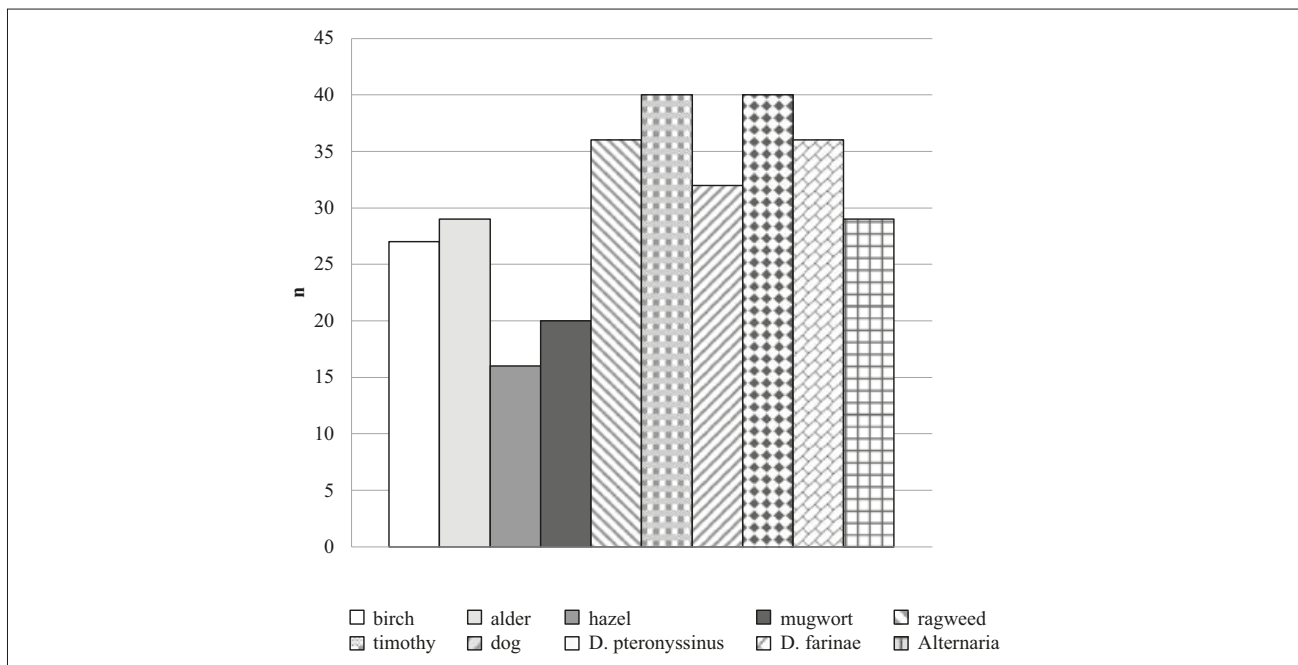


Fig. 1. Number of positive subjects (n) to aeroallergens, defined by skin tests

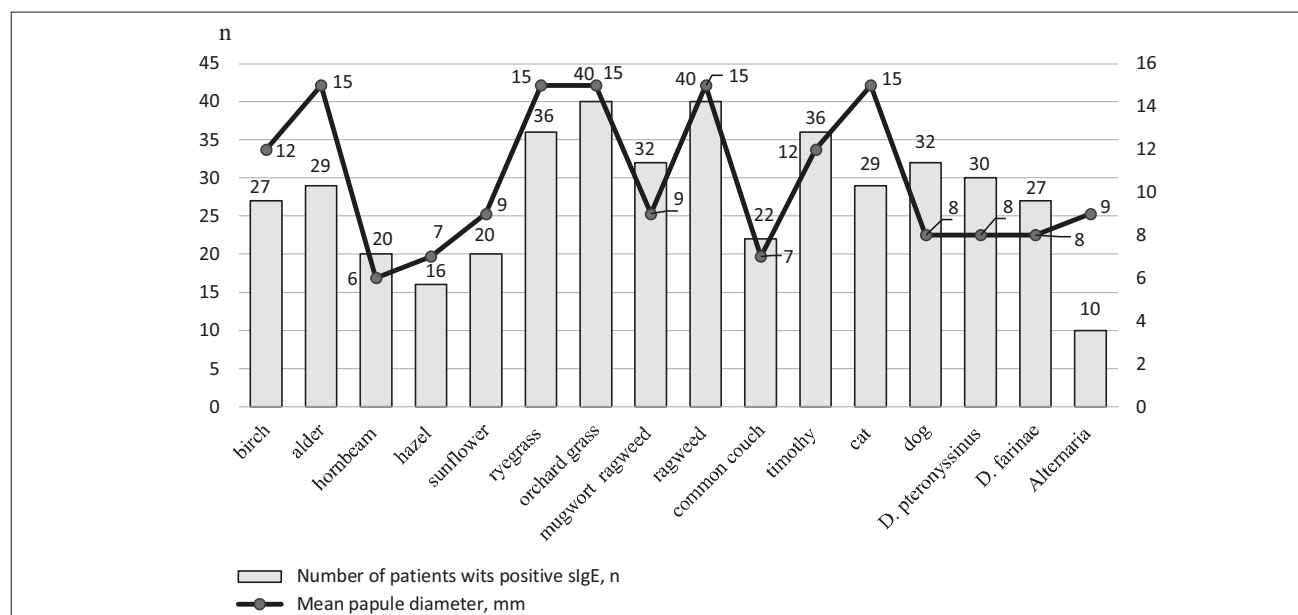


Fig. 2. Number of skin prick test positive subjects (n) with mean papule diameter (mm)

Excel editor. The relationship between clinical parameters was determined using the odds ratio (OR) with a 95% confidence interval (CI).

The research was conducted on the basis of the World Medical Association Declaration of Helsinki and the rules of good clinical practice (IGHGCP), voluntary participation, and informing patients about the nature of the research.

Results of the study and discussion

33 (33.7%) patients with asthma underwent skin testing when the diagnosis of asthma was made or within a year, according to the data anamnesis.

And 65 (66%) did not undergo an examination for the presence of allergies for more than a year after the diagnosis. Such data emphasize the fact that allergy diagnosis in asthma, a disease that largely depends on allergies, is delayed. To a large extent, this can be related to the insufficient education of primary care physicians and the difficult accessibility and cost of testing for the population.

Among 98 patients, skin testing with aeroallergens showed a positive result in 76 (77.6%) people: 18 (18.4%) showed positive skin testing for at least one allergen, and

58 (59.2%) had polysensitization. Individuals with polysensitized phenotypes usually have frequent multimorbidity, with severe and persistent course of disease.

Among those sensitized patients, 54 (55.1%) were sensitive to pollen allergens, 27 (27.6%) people had a positive test for birch, 29 (29.6%) people to alder, 16 (16.3%) people to hazel, hornbeam — 20 (20.4%), ryegrass — 36 (36.7%), orchard grass — 40 (40.8%), ragweed — 40 (40.8%), mugwort — 32 (32.6%), sunflower — 20 (20.4%), common couch — 22 (22.4%), timothy grass — 36 (36.7%). A positive skin test for cat allergen was found in 29 (29.6%), dog — 32 (32.6%), *Dermatophagoides pteronyssinus* — 30 (30.6%), *Dermatophagoides farinae* — 27 (27.6%), *Alternaria* — 10 (10.2%). The data on skin test results are presented in Figure 1.

As can be seen from Figure 2, the largest positive reactions to allergens were found on pollen: birch, alder, ryegrass, orchard grass, and ragweed. Although this analysis has certain drawbacks: the diameter of the papule may depend on the medication, the season of testing, and other individual factors. The prevalence of sensitization and allergy to various allergens depends on the population and differs in various literary sources. In particular, according to many data, the first place among common allergens belongs to timothy, birch, cat allergens [2,4,8,12]. Such data largely depend on the cohort of the surveyed: urban or rural population; the age of the examinees; geographical latitude and spectrum of plants; level of environmental pollution.

Children with moderate and severe asthma significantly more often had year-round AR than

seasonal AR (OR=3.3, CI 1.1650–9.6568, p=0.02). In this group, we did not find an association of polysensitization with a more severe asthma phenotype (OR=1.6, CI 0.5153 to 5.4584, p=0.39). Although literary data demonstrate such a connection of moderate to severe, exacerbation-prone asthma: the average number of inhalant sensitizations for school-age children was 14 positive results out of 22 tested allergens (68%) [13].

In total, 12 (12.2%) patients showed clinically not significant sensitization to allergens, and patients reported no symptoms after contact with allergens. In particular, 7 (7.1%) patients had sensitization to cats and/or dogs and had no symptoms.

Conclusions

Skin prick testing is an available research method that provides improvement in asthma care, gives opportunities for allergic trigger control, and defines which patients with asthma would be candidates for ASIT. The most important inhalant allergens are pollen (trees, grasses, and weeds), house dust mites, mold, and animal dander.

Yet most children with asthma do not receive ASIT because of the lack of awareness at the primary care level, lack of testing for aeroallergen sensitization, and financial inaccessibility of examination and treatment in Ukraine.

No conflict of interests was declared by the authors.

Acknowledgments. We thank the patients and their parents for participating in the study and for their cooperation.

REFERENCES/ЛІТЕРАТУРА

- Agache I, Ryan D, Rodriguez MR, Yusuf O, Angier E, Jutel M. (2013). Allergy management in primary care across European countries — actual status. *Allergy*. 68(7): 836–843. <https://doi.org/10.1111/all.12150>.
- Antypkin YuH, Volosovets OP. (2020). Air pollution and health of children in Ukraine. *Ukrainian Journal of Perinatology and Pediatrics*. 3(83): 31–39. [Антипкін ЮГ, Волосовець ОП. (2020). Забруднення повітря та стан здоров'я дитячого населення України. *Український журнал Перинатологія і Педіатрія*. 3(83): 31–39]. doi: 10.15574/PP.2020.83.31.
- Antypkin YuH, Volosovets OP, Maidannik VG, Berezenko VS, Moiseenko RA, Vygovskaya OV et al. (2018). Child health status — the future of the country (part 2). *Child's health*. 13(2): 142–152. [Антипкін ЮГ, Волосовець ОП, Майданник ВГ, Березенко ВС, Моїсеєнко РО, Виговська ОВ та інш. (2018). Стан здоров'я дитячого населення — майбут-
- не країни (частина 2). *Здоров'я дитини*. 13(2): 142–152]. doi: 10.22141/2224-0551.13.2.2018.129546.
- Casale TB, Pedersen S, Rodriguez Del Rio P, Liu AH, Demoly P, Price D. (2020). The Role of Aeroallergen Sensitization Testing in Asthma Management. *The journal of allergy and clinical immunology. In practice*. 8(8): 2526–2532. <https://doi.org/10.1016/j.jaip.2020.07.004>.
- Custovic A, Custovic D, Fontanella S. (2024). Understanding the heterogeneity of childhood allergic sensitization and its relationship with asthma. *Current opinion in allergy and clinical immunology*. 24(2): 79–87. <https://doi.org/10.1097/ACI.0000000000000967>.
- Custovic A, de Moira AP, Murray CS, Simpson A. (2023). Environmental influences on childhood asthma: Allergens. *Pediatric allergy and immunology : official publication*

- of the European Society of Pediatric Allergy and Immunology. 34(2): e13915. <https://doi.org/10.1111/pai.13915>.
7. GINA. (2023). Global Initiative for Asthma. Methodology. Fontana, Global Initiative for Asthma. URL: <https://ginasthma.org/wp-content/uploads/2023/07/GINA-2023-Pocket-Guide-WMS.pdf>.
 8. Martinez FD. (2019). Childhood Asthma Inception and Progression: Role of Microbial Exposures, Susceptibility to Viruses and Early Allergic Sensitization. *Immunology and allergy clinics of North America*. 39(2): 141–150. <https://doi.org/10.1016/j.iac.2018.12.001>.
 9. National Asthma Education and Prevention Program. (2007). Expert Panel Report 3 (EPR-3): Guidelines for the Diagnosis and Management of Asthma-Summary Report 2007. *The Journal of allergy and clinical immunology*. 120; 5 Suppl: S94–S138. <https://doi.org/10.1016/j.jaci.2007.09.043>.
 10. Niespodziana K, Borochova K, Pazderova P, Schleder T, Astafyeva N, Baranovskaya T et al. (2020). Toward personalization of asthma treatment according to trigger factors. *The Journal of allergy and clinical immunology*. 145(6): 1529–1534. <https://doi.org/10.1016/j.jaci.2020.02.001>.
 11. Oksel C, Custovic A. (2018). Development of allergic sensitization and its relevance to paediatric asthma. *Current opinion in allergy and clinical immunology*. 18(2): 109–116. <https://doi.org/10.1097/ACI.0000000000000430>.
 12. Papadopoulos NG, Miligkos M, Xerapadaki P. (2022). A Current Perspective of Allergic Asthma: From Mechanisms to Management. *Handbook of experimental pharmacology*. 268: 69–93. https://doi.org/10.1007/164_2021_483.
 13. Pongracic JA, Krouse RZ, Babineau DC, Zoratti EM, Cohen RT, Wood RA et al. (2016). Distinguishing characteristics of difficult-to-control asthma in inner-city children and adolescents. *The Journal of allergy and clinical immunology*. 138(4): 1030–1041. <https://doi.org/10.1016/j.jaci.2016.06.059>.
 14. Silverwood RJ, Rutter CE, Mitchell EA, Asher MI, Garcia-Marcos L, Strachan DP et al. (2019). Are environmental risk factors for current wheeze in the International Study of Asthma and Allergies in Childhood (ISAAC) phase three due to reverse causation? *Clinical and experimental allergy : journal of the British Society for Allergy and Clinical Immunology*. 49(4): 430–441. <https://doi.org/10.1111/cea.13325>.
 15. Volosovets O, Bolbot Y, Beketova G, Berezenko V, Umanets T, Rechkina O et al. (2021). Allergic march in children of Ukraine. *Medicni Perspektivi*. 26(4): 181–188. [Волосовець ОП, Більбот ЮК, Бекетова ГВ, Березенко ВС, Уманець ТР, Речкіна ОО та інш. (2021). Алергічний марш у дітей України. Медичні перспективи. 26(4): 181–188]. <https://doi.org/10.26641/2307-0404.2021.4.248227>.
 16. Volosovets OP, Bolbot YK, Kryvopustov SP, Mozyrska OV, Kryvopustova MV, Prokhorova MP, Kupkina AV. (2020). Bronchial asthma in children of Ukraine: medical and environmental parallels of morbidity and prevalence. *Medicni Perspektivi*. 25(3): 184–191. [Волосовець ОП, Більбот ЮК, Кривоустов СП, Мозирська ОВ, Кривоустова МВ, Прохорова МП, Купкіна АВ. (2020). Бронхіальна астма у дітей України: медико-екологічні паралелі захворюваності та поширеності. Медичні перспективи. 25(3): 184–191]. <https://doi.org/10.26641/2307-0404.2020.3.214861>.
 17. Volosovets OP, Kryvopustov SP, Mozyrska OV, Skvarkaya OO, Saltanova SD, Yemets OV, Karulina YuV. (2018). Post-chornobyl trends in the prevalence of diseases and incidence of the children's population in Ukraine. *World of medicine and biology*. 2(64): 15–24. doi:10.26724/2079-8334-2018-2-64-15-24.
 18. Volosovets OP, Kryvopustov SP, Volosovets TM, Abaturov OE, Kryuchko TO. (2019). Changes in health status of child population of Ukraine after Chernobyl catastrophe. *Wiadomości Lekarskie*. LXXII(10): 1974–1976. doi: <https://doi.org/10.36740/WLek201910123>.
 19. Volosovets O, Vyhovska O, Kryvopustov S, Mozyrska O, Yemets O, Volosovets A, Feleszko W. (2023). Problems of providing medical care to children of Ukraine as a result of russian aggression. *Child's health*. 18(3): 157–161. <https://doi.org/10.22141/2224-0551.18.3.2023.1578>.

Відомості про авторів:

Мозирська Олена Вікторівна — к.мед.н., доц. каф. педіатрії №2 НМУ ім. О.О. Богомольця. Адреса: м. Київ, вул. Алішера-Навої, 3. <https://orcid.org/0000-0001-9936-8304>.

Соколова Яна Вікторівна — лікар-педіатр КМДКЛ № 2. Адреса: м. Київ, вул. Алішера-Навої, 3. Стаття надійшла до редакції 02.06.2024 р., прийнята до друку 15.10.2024 р.