The importance of specific IgE antibodies in epidemiology of allergic rhinitis and asthma – the Epidemiology of Allergic Diseases in Poland (ECAP) survey: part one. Influence of allergy risk factors on concentration of specific IgE antibodies in serum

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Abstract

Introduction: Specific immunoglobulins E (sIgE) are important parameters to estimate severity of allergic diseases.

Aim: To determine the influence of allergy risk factors on the concentration of specific IgE antibodies in serum.

Material and methods: The concentration of sIgE antibodies against allergens Dermatophagoides pteronyssinus, cat dander, timothy grass, Alternaria alternata were determined in serum of 4077 respondents randomly selected from 9 regions (ECAP study). The positive results of sIgE (≥ 0.35 IU/ml) were correlated to answers in questionnaires ECRHSII and ISAAC.

Results: sIgE are more frequently detected in respondents declaring an allergic disease of a father than of a mother (D. pteronyssinus p < 0.05, A. alternata p < 0.01). An early beginning to attend school, kindergarten or nursery increases the frequency of sIgE detection (p < 0.05). If the number of children who slept in the same room as a respondent before the respondent was five years old, was lower, sIgE antibodies are more frequently detected (p < 0.05). Relating to: 1) all 4 allergens, sIgE are the most frequently detected in respondents declaring absence of a carpet/rug at home (p < 0.05), 2) cat dander – less frequently detected in respondents who have a cat at home at the age of 1–4 years (p < 0.005).

Conclusions: Carpets/rugs capture a portion of mite, epidermal, mould allergens located at home, wherefore absence of a carpet or rug causes greater stimulation of an immune system by these allergens; as a consequence, IgE antibodies are the most frequently detected in respondents. Household contacts of the respondents, at the age of 1–4 years, with cats induced partial immune tolerance to cat dander.

Key words: specific immunoglobulins E, allergy, epidemiology, ECAP

Introduction

Scientific studies accomplished within the last 20 years demonstrate that allergy and asthma are the most rapidly proliferating diseases in children's population, and affect even more than 30% of infants in developed countries. Allergy and asthma are currently diseases of affluence, a major prevalence and morbidity rate for people less than 30 years old, suffering from non-infectious chronic diseases [1–4]. Epidemiological analysis shows that inhabitants of urban areas redevelop allergies more frequently than inhabitants of rural areas [5]. In patients with allergic rhinitis, the risk of asthma is several times higher [2]. In turn, asthma significantly impacts on the quality of patients' life and may socially exclude some of them. Therefore, it is remarkably important to find risk factors for periodic rhinitis, chronic rhinitis and asthma. Numerous studies were performed as part of the Epidemiology of Allergic Diseases in Poland (ECAP) survey, proving epidemiological significance.
of these diseases and great diversity of allergy risk factors [6–9]. The obtained results have evidenced, among others, meaningfully different sensitivity to common inhalation allergens, particularly comparing inhabitants of urban areas to inhabitants of villages. It was necessary to supplement results of this survey by determination of specific IgE in respondents’ serum. Determination of specific IgE is the most modern and the most reliable method to evaluate allergic hypersensitivity [10, 11]. The present study is a continuation of the ECAP survey, epidemiological studies of allergy and asthma, performed in 9 regions of Poland [12].

Aim

Aim of the study was to determine the influence of allergy risk factors on the concentration of specific IgE antibodies in serum. Additionally, aims of the study described in this article were: a) to evaluate the prevalence of sensitivity to the most important inhalation allergens (Dermatophagoides pteronyssinus, cat dander, timothy grass, Alternaria alternata) in the population of Poland, b) to evaluate the influence of sensitivity to aeroallergens on the prevalence of allergic rhinitis and asthma.

Material and methods

Twenty two thousand seven hundred and three respondents were randomly selected, using their personal identity numbers, from 8 cities with a population in excess of 150 000 and one rural region. The survey was based on international standards ECRHS II, using connected questionnaires ECRHS II and ISAAC [12]. Four thousand seven hundred eighty-three respondents were randomly selected, using their personal belongings of the study, performed in 9 regions of Poland [12].

Table 1. Number (percentage) of respondents with sIgE concentration \( \geq 0.35 \) IU/ml (classes 1–6)

| Relatives with an allergic disease | Respondents’ sIgE against |
|-----------------------------------|--------------------------|
| D. pteronyssinus (d1)             | Cat dander (e1)          |
|                                  | Timothy grass (g6)      |
|                                  | A. alternata (m6)       |
| **N (100%)**                     |                          |
| Mother                           | 47 (15.3%)               |
|                                  | 23 (7.5%)                |
|                                  | 44 (14.3%)               |
|                                  | 8 (2.6%)                 |
| Father                           | 44 (24.3%)               |
|                                  | 18 (9.9%)                |
|                                  | 31 (17.1%)               |
|                                  | 14 (7.7%)                |
| Sibling(s)                       | 76 (16.7%)               |
|                                  | 39 (8.6%)                |
|                                  | 78 (17.1%)               |
|                                  | 15 (3.3%)                |
| Paternal grandparents            | 11 (12.6%)               |
|                                  | 2 (2.3%)                 |
|                                  | 10 (11.5%)               |
|                                  | 2 (2.3%)                 |
| Maternal grandparents            | 6 (13.0%)                |
|                                  | 2 (4.4%)                 |
|                                  | 2 (4.4%)                 |
|                                  | 1 (2.2%)                 |
| Anyone at all                    | 267 (17.9%)              |
|                                  | 119 (8.0%)               |
|                                  | 240 (16.1%)              |
|                                  | 72 (4.8%)                |
| No one                           | 327 (12.8%)              |
|                                  | 120 (4.7%)               |
|                                  | 283 (11.1%)              |
|                                  | 80 (3.1%)                |

N includes respondents also declaring an allergic disease in other relatives(s): mother 592, father 371, sibling(s) 720, paternal grandparents 222, maternal grandparents 122.
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Statistical analysis

The aim of the statistical analysis was to compare proportions of people with a high level of immunoglobulin in two groups. The classical approximate test for comparison of two proportions was applied [14]. If calculated p-value was smaller than 0.05, the statistically

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Figure 1. Influence on detection of specific IgE antibodies (classes 1–6) against Dermatophagoides pteronyssinus, cat dander, timothy grass, Alternaria alternata in respondents’ serum

|                | OR    | Lower CL | Upper CL |
|----------------|-------|----------|----------|
| **Anyone at all** (Table 1) D. pteronyssinus (d1) | 1.4925 | 1.3164 | 1.6686 |
|                | Cat dander (e1) | 1.7642 | 1.5021 | 2.0263 |
|                | Timothy grass (g6) | 1.5475 | 1.3622 | 1.7328 |
|                | A. alternata (m6) | 1.5807 | 1.2557 | 1.9057 |
| **2–5 years** (Table 2 A) D. pteronyssinus (d1) | 1.1381 | 0.9575 | 1.3187 |
|                | Cat dander (e1) | 1.7071 | 1.4168 | 1.9974 |
|                | Timothy grass (g6) | 1.4235 | 1.228 | 1.619 |
|                | A. alternata (m6) | 2.1739 | 1.7945 | 2.5533 |
| **6 years** D. pteronyssinus (d1) | 0.9285 | 0.7364 | 1.1206 |
|                | Cat dander (e1) | 0.6993 | 0.3916 | 1.007 |
|                | Timothy grass (g6) | 0.8431 | 0.6373 | 1.0489 |
|                | A. alternata (m6) | 0.6009 | 0.205 | 0.9968 |
| **7 years** D. pteronyssinus (d1) | 0.5586 | 0.1016 | 1.0156 |
|                | Cat dander (e1) | 0.1991 | −0.9475 | 1.3457 |
|                | Timothy grass (g6) | 0.4442 | −0.0872 | 0.9756 |
|                | A. alternata (m6) | 0.2092 | −1.1922 | 1.6106 |
| **No children** (Table 2 B) Cat dander (e1) | 1.287 | 1.0225 | 1.5515 |
|                | Timothy grass (g6) | 1.3207 | 1.1348 | 1.5066 |
|                | A. alternata (m6) | 1.497 | 1.1723 | 1.8217 |
| **One child** Cat dander (e1) | 1.0343 | 0.7701 | 1.2985 |
|                | Timothy grass (g6) | 0.8691 | 0.6818 | 1.0564 |
|                | A. alternata (m6) | 0.8606 | 0.5301 | 1.1911 |
| **More than two children** Cat dander (e1) | 0.5763 | 0.1672 | 0.9854 |
|                | Timothy grass (g6) | 0.7889 | 0.5335 | 1.0443 |
|                | A. alternata (m6) | 0.6071 | 0.1083 | 1.1059 |
| **Absence of a carpet or rug** (Table 3 A) D. pteronyssinus (d1) | 1.3924 | 1.1899 | 1.5949 |
|                | Cat dander (e1) | 1.4003 | 1.1019 | 1.6987 |
|                | A. alternata (m6) | 1.5382 | 1.1767 | 1.8997 |
| **Absence of a carpet or rug** (Table 3 B) D. pteronyssinus (d1) | 1.4479 | 1.2516 | 1.6442 |
|                | Cat dander (e1) | 1.4448 | 1.1545 | 1.7351 |
|                | A. alternata (m6) | 1.4774 | 1.121 | 1.8338 |
| **Yes** (Table 4) Cat dander (e1) | 1.4503 | 1.1288 | 1.7718 |
| **No cat, at least 1–4 years** Cat dander (e1) | 1.3988 | 1.121 | 1.6766 |

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Did your mother, father or carer smoke cigarettes when you were a child?

Did your mother or carer smoke in the first year of your life?

The study was approved by the Institutional Bioethics Committee.
significant difference between investigated proportions was recognised. Otherwise, the fractions of people with a high level of immunoglobulin in investigated groups may be treated as similar. Calculations were performed using the statistical package Statistica (Statistica, Tulsa, Oklahoma, US).

Results

IgE antibodies are more frequently detected in respondents declaring an allergic disease of immediate relatives (D. pteronyssinus \( p < 0.001 \), cat dander \( p < 0.001 \), timothy grass \( p < 0.001 \), A. alternata \( p < 0.01 \)), the allergic disease of a father is of considerable importance ("allergic disease of a father" versus "allergic disease of a mother": D. pteronyssinus \( p < 0.05 \), A. alternata \( p < 0.01 \) (Table 1, Figure 1). There was no statistically significant difference between influence of the allergic disease of maternal grandparents and influence of the allergic disease of paternal grandparents.

An early beginning to attend school, kindergarten or nursery increases frequency of IgE antibodies detection in respondents (Table 2, Figure 1). Relating to all allergens, numerous statistically significant differences were identified (\( p < 0.05 \) to \( p < 0.001 \)), comparing groups aged 2–5 years to a group aged 6 years, groups aged 2–5 years to a group aged 7 years, and a group aged 6 years to a group aged 7 years.

Table 2. Number (percentage) of respondents with sIgE concentration ≥ 0.35 IU/ml (classes 1–6). A – Age of first attending school, kindergarten or nursery, B – number of other children who regularly slept in the same room as a respondent before the respondent was 5 years old

| Variable | Respondents’ sIgE against |
|----------|---------------------------|
|          | D. pteronyssinus (d1) | Cat dander (e1) | Timothy grass (g6) | A. alternata (m6) | N (100%) |
| A        |                          |                |                      |                    |
| 1 year   | 24 (17.9%)               | 7 (5.2%)       | 12 (9.0%)            | 2 (1.5%)           | 134      |
| 2 years  | 45 (17.6%)               | 16 (6.3%)      | 32 (12.5%)           | 13 (5.1%)          | 256      |
| 3 years  | 154 (14.4%)              | 69 (6.4%)      | 153 (14.3%)          | 50 (4.7%)          | 1073     |
| 4 years  | 90 (14.3%)               | 51 (8.1%)      | 94 (14.9%)           | 34 (5.4%)          | 630      |
| 5 years  | 82 (17.2%)               | 33 (6.9%)      | 72 (15.1%)           | 19 (4.0%)          | 477      |
| 6 years  | 171 (14.0%)              | 56 (4.6%)      | 142 (11.6%)          | 32 (2.6%)          | 1225     |
| 7 years  | 21 (8.9%)                | 3 (1.3%)       | 15 (6.4%)            | 2 (0.9%)           | 235      |
| B        |                          |                |                      |                    |
| 0        | 223 (14.3%)              | 105 (6.7%)     | 231 (14.8%)          | 73 (4.7%)          | 1560     |
| 1        | 259 (14.7%)              | 105 (6.0%)     | 211 (12.0%)          | 61 (3.5%)          | 1762     |
| 2+       | 109 (15.1%)              | 27 (3.7%)      | 78 (10.8%)           | 18 (2.5%)          | 722      |

Table 3. Number (percentage) of respondents with sIgE concentration ≥ 0.35 IU/ml (classes 1–6). A – How old is the oldest carpet or rug in the room which a respondent uses most at home during the day, B – how old is the oldest carpet or rug in the room where a respondent sleeps

| Variable | Respondents’ sIgE against |
|----------|---------------------------|
|          | D. pteronyssinus (d1) | Cat dander (e1) | A. alternata (m6) | N (100%) |
| A        |                          |                |                    |
| Absence of a carpet or rug | 152 (18.2%) | 63 (7.5%) | 43 (5.2%) | 835 |
| Less than 1 year | 45 (13.4%) | 11 (3.3%) | 16 (4.8%) | 337 |
| 1–5 years | 193 (13.1%) | 85 (5.8%) | 50 (3.4%) | 1470 |
| More than 5 years | 203 (14.6%) | 80 (5.7%) | 43 (3.1%) | 1394 |
| B        |                          |                |                    |
| Absence of a carpet or rug | 168 (18.6%) | 69 (7.6%) | 45 (5.0%) | 905 |
| Less than 1 year | 43 (12.4%) | 13 (3.8%) | 15 (4.3%) | 346 |
| 1–5 years | 203 (13.6%) | 91 (6.1%) | 53 (3.6%) | 1493 |
| More than 5 years | 180 (13.9%) | 65 (5.0%) | 39 (3.0%) | 905 |
If the number of other children, who regularly slept in the same room as a respondent before the respondent was 5 years old, is lower, IgE antibodies are more frequently detected in respondents (Table 2, Figure 1). Relating to allergens of cat dander, timothy grass, and A. alternata, numerous statistically significant differences were identified (\( p < 0.05 \) to \( p < 0.005 \)), comparing groups 0, 1, 2+, especially group “0” to “2+”.

Relating to allergens of D. pteronyssinus, cat dander, and A. alternata, IgE antibodies are the most frequently detected in respondents declaring absence of a carpet or rug in the room which the respondent uses most at home during the day or in the room where the respondent sleeps (Table 3, Figure 1). Numerous statistically significant differences were identified (\( p < 0.05 \) to \( p = 0.001 \)), comparing a group “absence of a carpet or rug” to every remaining group. Very similar values were obtained after excluding, from the statistical analysis, respondents declaring they have ever had asthma. Thus, the observed relationship does not result from asthmatics conforming to doctor’s orders, and removing carpets and rugs from home.

Relating to an allergen of cat dander, IgE antibodies are more frequently detected in respondents keeping a cat (\( p < 0.05 \)), but less frequently detected in respondents declaring there was a cat at home when they were aged 1–4 years (“cat at home, at least, when respondents were aged 1–4 years” versus “no cat at home when respondents were aged 1–4 years” \( p < 0.005 \)) (Table 4, Figure 1).

IgE antibodies are more frequently detected in respondents living in a city, most of the time when they were under the age of 5 years (“rural” versus “urban”: cat dander \( p < 0.001 \), timothy grass \( p < 0.001 \), A. alternata \( p < 0.001 \)) (Table 5, Figure 1).

### Discussion
Numerous studies were performed as part of the ECAP survey, proving epidemiological significance of these diseases and great diversity of allergy risk factors. It was necessary to supplement results of this survey by determination of specific IgE in respondents’ serum. The aim of the study described in this article was to determine influence of allergy risk factors on the concentration of specific IgE antibodies in serum.

IgE antibodies are more frequently detected in respondents declaring an allergic disease of a father than in respondents declaring an allergic disease of a mother. This results, probably, from immune tolerance to some of potential allergens, as a consequence of interaction with the allergic mother’s immune system, in the course of a foetal period. In a study by Hensley Alford et al., father’s allergic disease history, particularly asthma history, was more strongly related to paediatric outcomes than mother’s history [15]; in a study by Anderson et al., a history of paternal asthma, and to some degree allergy, appears to confer an increased risk of allergic sensitization in preschool children, similar maternal histories are not significantly associated with these developments [16]. On the other hand, in a study by Mandhane et al., maternal atopy but not paternal atopy was significantly associated with asthma of male children [17], while in a meta-analysis by Lim et al., maternal asthma increases the offspring disease risk to a greater extent than paternal disease [18]. Moreover, in a study by Arshad et al., maternal asthma was associated with asthma in girls but not in boys, whereas paternal asthma was associated with asthma in boys but not in girls [19]. In a study by Westman et al., parental allergy-related disease may be an important risk factor for nonallergic rhinitis as well as allergic rhinitis, and the risk is comparable for maternal and paternal allergy [20]. In a study by Fuertes et al., parental allergic diseases both pose risks to childhood

### Table 4. Number (percentage) of respondents with sIgE concentration \( \geq 0.35 \) IU/ml (classes 1–6). A – Respondent keeps a cat, B – age of a respondent when a cat was at home (the first year of life, 1–4 years, 4–15 years)

| Variable | Cat dander (e1) | N (100%) |
|----------|----------------|----------|
| A        |                |          |
| Yes      | 51 (7.8%)      | 653      |
| No       | 188 (5.5%)     | 3403     |
| B        |                |          |
| Never    | 183 (6.3%)     | 2924     |
| First year of life | 3 (2.5%) | 121 |
| 0–4 years | 5 (11.4%) | 44 |
| 0–15 years | 8 (3.1%) | 255 |
| 1–4 years | 6 (3.2%) | 187 |
| 1–15 years | 2 (1.5%) | 130 |
| 4–15 years | 31 (7.9%) | 395 |

### Table 5. Number (percentage) of respondents with sIgE concentration \( \geq 0.35 \) IU/ml (classes 1–6). Relating to the place where a respondent, aged less than 5 years, lived most of the time

| Place     | Respondents’ sIgE against |
|-----------|----------------------------|
|           | D. pteronyssinus (d1) | Cat dander (e1) | Timothy grass (g6) | A. alternata (m6) | N (100%) |
| Rural     | 99 (13.3%) | 21 (2.8%) | 53 (7.1%) | 10 (1.4%) | 743 |
| Urban     | 494 (14.9%) | 218 (6.6%) | 470 (14.2%) | 142 (4.3%) | 3310 |
allergic disease in the offspring, especially for asthma [21]. An early beginning to attend school, kindergarten or nursery increases the frequency of IgE antibodies detection in respondents. This results, most likely, from a greater diversity of allergens, which more strongly stimulates an immune system to allergic reactions, in humans exposed to these allergens in the first year of their life. The school environment can be an important size of exposure to indoor allergens detected in settled school dust [22]. In a study by Sporik et al., exposure in early childhood to house dust mite allergens is an important determinant of the subsequent development of asthma [23], while in a study by Cole Johnson et al., dust mite exposure in early childhood is associated with a higher risk of sensitization in the presence of a positive parental history of atopic disease [24], in infants and preschool children. Authors of ARIA 2010 suggest multifaceted interventions to reduce early life exposure to house dust mite [25]. On the other hand, regular sleeping in the same room as other children, before a respondent was 5 years old, decreases the frequency of IgE antibodies detection in respondents. The co-sleeping leads to more frequent infections, which presumably decreases probability of prospective allergic reactions. Numerous studies prove that a higher number of siblings protects against the development of allergy and asthma [26]. Carpets and rugs capture a portion of mite, epidermal, mould allergens located at home, wherefore absence of a carpet or rug causes greater stimulation of an immune system by these allergens; as a consequence, IgE antibodies are the most frequently detected in respondents. House dust mite exposure is dose-dependently related to a higher risk of sensitization to mites [27]. IgE antibodies against cat dander are more frequently detected in respondents keeping a cat, nonetheless household contacts of the respondents, at the age of 1 to 4 years, with cats induced partial immune tolerance to this allergen. Celedon et al. showed that in children who did not have a maternal history of asthma, exposure to Fel d 1 cat antigen at the age of 2 to 3 months was associated with a reduced risk of wheezing, but in children with a maternal history of asthma, similar exposure was associated with an increased risk of wheezing, and the risk associated with cat exposure became greater with increasing duration of follow-up [28]. In infants and preschool children, authors of ARIA 2010 suggest no special avoidance of exposure to pets at home [25].

Conclusions

IgE antibodies are more frequently detected in respondents declaring an allergic disease of a father than in respondents declaring an allergic disease of a mother. An early beginning to attend school, kindergarten or nursery increases frequency of IgE antibodies detection in respondents. Regular sleeping in the same room as other children, before a respondent was 5 years old, decreases frequency of IgE antibodies detection in respondents. Carpets and rugs capture a portion of mite, epidermal, mould allergens located at home, wherefore absence of a carpet or rug causes greater stimulation of an immune system by these allergens; as a consequence, IgE antibodies are the most frequently detected in respondents. IgE antibodies against cat dander are more frequently detected in respondents keeping a cat, nonetheless household contacts of the respondents, at the age of 1 to 4 years, with cats induced partial immune tolerance to this allergen.

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Conflict of interest

The authors declare no conflict of interest.

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