Immediate Hypersensitivity to Common Food Allergens: An Investigation on Food Sensitization in Respiratory Allergic Patients of Calcutta, India

Jyotshna Mandal,1 Mahasweta Das,2 Indrani Roy,3 Soma Chatterjee,3 Nimai Chandra Barui,2 and Swati Gupta-Bhattacharya1

Background: Food allergy may be defined as an immunoglobulin E–mediated immune response to food proteins. Such studies have previously not been done in Calcutta, India. The present study was therefore undertaken to record the sensitivity to commonly consumed foods in patients with allergic rhinitis and asthma.

Materials and Methods: A survey of 800 patients (410 males and 390 females) reporting to the Allergy Unit of the Institute of Child Health, Calcutta, were selected for the study conducted from May 2006 to April 2007. Respiratory allergic patients in the age group of 5 to 60 years were evaluated using a standard questionnaire, and skin prick test was performed using common food and aeroallergens.

Results/Conclusions: Out of the 684 patients with a history of food allergy, 338, are in the age group 16 to 40 years, 192 of them were in the age group 16 to 40 years, 192 of them were in the age group 41 to 60 years, and 154 were in the age group 5 to 15 years. Most of the patients with food allergy had asthma (65.05%), rhinitis and asthma (20.03%), and skin allergies (4.97%), such as itching, eczema, and urticaria. The foodstuffs that were found to elicit symptoms of hypersensitivity were egg, milk, wheat, pulses, vegetables, fishes, and fruits.

Food allergy is a major health problem affecting 6% of young children and 3% to 4% of adults.1 The incidence of perceived food hypersensitivity varies (1.4%-1.9%) largely across different countries probably because of differences in the diagnostic criteria, study design and population variation, and so on.2 In young children, the common causal food allergens are cow’s milk, egg, peanut, wheat, soy, tree nuts, fish, and shellfish, whereas adults mostly have allergies to shellfish, peanuts, tree nuts, and fish.3 The order of importance of specific allergens varies in different countries, reflecting a possible interaction of genetic factors, cultural and dietary habits, and exposure to new allergenic products early in life.4 Several food allergies to fruits and nuts have become increasingly common and represent a growing clinical problem.5 Studies have shown that food allergy in adolescent and adult individuals develops because of an allergic sensitization or cross reaction to inhalant pollen allergens.6,7

In India, there has been an increase in various allergic diseases from 10% to 30% in the last 4 decades.8 Many asthma and rhinitis patients may be having food allergy, but only a few studies on food hypersensitivity have been carried out so far in India.9-11 These studies indicate that common food allergens such as egg, milk, cereals, and legumes induce immunoglobulin E–mediated reactions in children as well as in the adult population.12

The present study has been designed to determine the incidence of allergic responses toward common food allergens among the respiratory allergy population of Calcutta metropolis, India.

MATERIALS AND METHODS

Preparation of Extracts

Various food allergens were defatted in diethyl ether at 4°C. The extraction was carried out by continuous stirring for 8 hours at 4°C in 1:20 wt/vol phosphate buffered saline ([PBS] 0.1 M sodium phosphate, pH 7.2). After centrifugation at 12,000g, the clear supernatant was dialyzed and passed through a 0.22-μm Millipore filter (Millipore Corp, Bedford, Mass). The filtrate was then lyophilized and stored at −70°C in aliquots of known volume in sterile vials.

Study Subjects

A total of 800 patients (410 males and 390 females) aged 5 to 60 years reporting to the Allergy Unit of the Institute of Child Health, Calcutta, were screened for food hypersensitivity using a standard questionnaire and skin prick test (SPT). The study was conducted from May 2006 to April 2007, and it is composed of patients with a history of bronchial asthma, allergic rhinitis, and urticaria, either alone or in...
different combinations. The exclusion criteria were perennial or severe asthma, pregnancy or lactation, malignancy, or other severe systemic diseases during skin testing. To avoid masking of severe symptoms, corticosteroids and antihistamines were prohibited. A detailed case history of the subjects was taken based on a structured questionnaire containing information regarding age, sex, religion, occupation, family history, type of diet (vegetarian or nonvegetarian), food habit, smoking habit, onset, duration, and the present status of the symptoms. Patients referred for allergy testing were asked whether they had hypersensitivity symptoms from any foodstuffs listed. Those who answered yes were asked to fill in the questionnaire in which 24 foods were listed along with the other previously mentioned details. The patients were asked specific questions to which they replied whether they had slight, moderate, or severe symptoms, or reported not having eaten that foodstuff. They were also asked to state the symptoms they got from that food hypersensitivity. Food hypersensitivity was ruled out if the foodstuffs had given rise to symptoms only once and did not give rise to any complication if they were repeatedly ingested afterward. Sixty healthy individuals from the city belonging to the same age group were also selected to act as control subjects (confirmed by negative skin reaction). Both male and female patients were categorized into 3 groups: group A (aged 5–15 years), group B (aged 16–40 years), and group C (aged 41–60 years). The study was approved by the ethics

**TABLE 1. Distribution of Patients as Per Symptom and History of Respiratory Allergy**

| Disease                  | Total Patients (n = 800; male-female ratio, 410:390) | Patients With a History of Food Allergy (n = 684; male-female ratio, 370:314) |
|--------------------------|-------------------------------------------------------|-------------------------------------------------------------------------------|
|                          | n         | %       | n         | %       |
| Bronchial asthma         | 145       | 18.12   | 123       | 17.98   |
| Allergic rhinitis        | 36        | 4.5     | 29        | 4.24    |
| Asthma with rhinitis     | 610       | 76.25   | 512       | 74.85   |
| Asthma with skin allergies | 12       | 1.5     | 9         | 1.32    |

**TABLE 2. Sensitivity to Different Food Allergens Among the Respiratory Allergic Patients in Various Age Groups**

| Allergens     | 5–15, n = 154 (M, 88; F, 66) | 16–40, n = 338 (M, 184; F, 154) | 41–60, n = 192 (M, 98; F, 94) |
|---------------|------------------------------|---------------------------------|-------------------------------|
|               | Total No. Patients Tested    | No. Patients Positive (%)       | No. Patients Showing >1 Level of Reaction (%) | Total No. Patients Tested | No. Patients Positive (%) | No. Patients Showing >1 Level of Reaction (%) | Total No. Patients Tested | No. Patients Positive (%) | No. Patients Showing >1 Level of Reaction (%) |
| Wheat (cereal)| 138                          | 30 (21.73)                      | —                              | 300                          | 100 (33.3)               | 8 (2.6)                          | 110                          | 26 (23.63)               | 4 (3.6)                          |
| Egg           | 150                          | 56 (37.3)                       | 12 (21.4)                      | 252                          | 78 (31)                  | 16 (63.4)                       | 96                          | 40 (41.6)                | 6 (62.5)                       |
| Fish          | 16                           | 6 (37.5)                        | —                              | 30                           | 10 (33.3)                | 1 (3.3)                         | 0                           | 0                          | —                              |
| T. ilisha     | 18                           | 8 (44.4)                        | 3 (16.6)                       | 26                           | 16 (61.5)                | 6 (23)                          | 12                          | 6 (50)                    | 7 (58.3)                       |
| Prawn         | 98                           | 6 (6.1)                         | 2                              | 144                          | 12 (8.3)                 | 2 (1.4)                         | 62                          | 0                          | —                              |
| Fruits        |                              |                                 |                                |                              |                        |                                 |                              |                            |                                 |
| Apple         | 24                           | 0                               | —                              | 32                           | 4 (12.5)                 | —                              | 80                          | 5 (6.25)                  | —                              |
| Banana        | 118                          | 48 (40.67)                      | 4 (3.38)                       | 138                          | 56 (40.57)               | 5 (3.6)                         | 74                          | 30 (40.54)               | —                              |
| Cucumber      | 30                           | 6 (20)                          | —                              | 86                           | 6 (6.97)                 | —                              | 66                          | 16 (24.24)               | —                              |
| Guava         | 22                           | 6 (27.27)                       | —                              | 36                           | 10 (27.7)                | —                              | 65                          | 15 (23.07)               | —                              |
| Papaya        | 76                           | 22 (28.95)                      | 2 (2.63)                       | 120                          | 40 (33.3)                | 2 (1.66)                        | 40                          | 0                          | 3 (7.5)                        |
| Pumpkin       | 64                           | 10 (15.63)                      | 1 (1.56)                       | 140                          | 30 (21.42)               | 2 (1.42)                        | 50                          | 9 (18)                    | 1 (2)                          |
| Grapes        | 16                           | 4 (25)                          | —                              | 68                           | 15 (22)                  | 2 (2.94)                        | 62                          | 10 (1.6)                  | —                              |
| Meat          |                              |                                 |                                |                              |                        |                                 |                              |                            |                                 |
| Beef          | 18                           | 4 (22.2)                        | —                              | 68                           | 14 (20.6)                | —                              | 0                           | 0                          | —                              |
| Chicken       | 78                           | 28 (35.9)                       | —                              | 98                           | 10 (10.20)               | 3 (3.06)                        | 40                          | 2 (5.0)                   | —                              |
| Mutton        | 24                           | 0                               | —                              | 40                           | 6 (15.0)                 | 1 (2.5)                         | 80                          | 10 (12.5)                | —                              |
| Milk          | 144                          | 36 (25)                         | 6 (4.16)                       | 304                          | 56 (18.42)               | 11 (3.6)                        | 106                         | 22 (20.75)               | 3 (1.88)                       |
| L. culinaris  | 96                           | 18 (18.75)                      | 1 (1.04)                       | 142                          | 16 (11.26)               | 8 (5.63)                        | 64                          | 14 (21.8)                | 4 (6.25)                       |
| P. mungo      | 22                           | 4 (18.1)                        | 1 (4.54)                       | 68                           | 8 (11.76)                | 5 (7.35)                        | 36                          | 12 (33.33)               | 2 (5.5)                        |
| Vegetables    |                              |                                 |                                |                              |                        |                                 |                              |                            |                                 |
| Brinjal       | 62                           | 28 (45.16)                      | 3 (4.83)                       | 152                          | 66 (43.42)               | 16 (10.52)                      | 40                          | 22 (55)                   | 3 (7.5)                        |
| Cabbage       | 44                           | 6 (13.6)                        | —                              | 104                          | 12 (11.53)               | 1 (0.96)                        | 66                          | 18 (27.27)               | —                              |
| Cauliflower   | 64                           | 16 (25)                         | 3 (4.6)                        | 152                          | 40 (26.3)                | —                              | 68                          | 26 (38.23)               | 2 (2.94)                       |
| Ladyfinger    | 104                          | 30 (28.8)                       | 2 (1.92)                       | 202                          | 72 (35.6)                | 4 (1.98)                        | 88                          | 36 (17.25)               | 1 (1.13)                       |
| Spinach       | 46                           | 18 (39.1)                       | 1 (2.17)                       | 84                           | 20 (23.8)                | 3 (3.57)                        | 50                          | 10 (20)                   | 1 (2)                          |
| Tomato        | 140                          | 34 (24.3)                       | 2 (1.43)                       | 258                          | 60 (23.25)               | 8 (3.1)                         | 78                          | 14 (17.95)               | —                              |

F indicates total number of females tested; M, total number of males tested.
committee of the hospital, and informed consents were obtained from the subjects before their participation.

Skin Prick Test

Skin prick test was performed with a panel of 24 commonly consumed foods—milk, egg, cereals (wheat), pulses (Lens culinaris, Phaseolus mungo), vegetables (brinjal, cabbage, cauliflower, ladyfinger, spinach, tomato), fruits (apple, banana, cucumber, guava, papaya, pumpkin, grapes), meat (beef, chicken, mutton), fish (Tenualosa ilisha [locally known as Hilsa], prawn, Labeo rohita [locally known as Rohu]), and 12 other common inhalant allergens (pollen and fungi, data not given)—on patients with symptoms suggestive of food allergy (history). Briefly, a drop (20 μL of each extract, 1:50 wt/vol in PBS [wt/vol in glycerinated PBS]) was placed on the forearm, and the skin was pricked with a 26G hypodermic needle.13 Histamine diphosphate (1 mg/mL) and PBS (1 mg/mL) were used as positive and negative controls, respectively. The reaction measurement was performed after 20 minutes. According to international guidelines, positivity was defined as a mean wheal diameter of 3 mm or more compared with the negative control.14 The reaction was graded from +1 to +3 level (+1, erythema 20 mm in diameter; +2, erythema >20 mm in diameter; +3, wheal and erythema) according to Stytis et al.15 The skin tests were graded after 20 minutes in comparison with the wheal diameter of the positive control.

RESULTS

Out of the total number of patients reporting to the clinic, 684 patients (85.5%) were reported to have a history of food allergy, most of them are in the age group 16 to 40 years (n = 338 [49.41%]), followed by 41 to 60 years (n = 192 [28.07%]). One hundred fifty-four (22.51%) belonged to the age group 5 to 15 years. It was observed that most of the patients with food allergy had asthma (17.98%), rhinitis and asthma (74.85%), rhinitis (4.24%), and a small number of patients had skin allergies (1.32%), such as itching, eczema, and urticaria (Table 1). The foodstuffs that were found to be able to have symptoms of hypersensitivity were egg, milk, wheat, pulses, vegetables, fishes, and fruits (Table 2).

In the age range 16 to 40 years (male-female ratio, 1:1.19), 33.3% were sensitive to prawn; brinjal, 43.42%; banana, 40.57%; ladyfinger, 35.6%; papaya, 33.3%; wheat, 33.3%; egg, 31%; cauliflower, 26.3%; spinach, 23.8%; beef, 20.6%; milk, 18.42%; lentil (P. mungo), 11.76%; L. culinaris, 11.26%; and cabbage, 11.53%. The age group 41 to 60 years (male-female ratio, 1:1.04) had skin reactivity to brinjal (55%), egg (41.6%), banana (40.54%), fish (38%), P. mungo (33.33%), cabbage (27.27%), wheat (23.63%), L. culinaris (21.8%), milk (20.75%), spinach (20%), and tomato (17.95%). Patients younger than 16 years were sensitized to brinjal (45.16%), prawn (44.4%), banana (40.67), spinach (39.1%), egg (37.3%), papaya (28.95%), ladyfinger (28.8%), cauliflower (25%), milk (25%), wheat (21.73%), L. culinaris (18.75%), P. mungo (18.1%), and cabbage (13.6%).

A list of all the allergens and the total number of patients positive to all these allergens are represented in Figure 1 and Table 3.

CONCLUSIONS

The present study was undertaken to identify food sensitization in patients aged 5 to 60 years with respiratory allergic symptoms, who reported to our allergy clinic in Calcutta. A total of 684 patients presented a history of allergy to different foods. Among them, 516 (75.44%) patients had a history of allergy to common food allergens such as egg, milk, cereals, and legumes that induce immunoglobulin E–mediated reactions. Among the common food allergens, allergy to egg was highest in the age group 41 to 60 years (41.6%); the frequency

FIGURE 1. Frequency of positive response to common food allergens among the respiratory allergy population in Calcutta, India.
of milk allergy was highest in the age group 5 to 15 years (25%), children being the largest consumers of milk; the percentage of hypersensitivity to cereals was highest among patients in their late teens and adults, that is, in the age group 16 to 40 years (33.3%).

These findings are in accordance with other reports on food hypersensitivity in children.4,16,17 Allergy to legumes (L. culinaris and P. mungo) was maximum in the age group 41 to 60 years (26%). Legume hypersensitivity on respiratory allergy patients has been similarly reported in India.9,12,18 It was observed that 315 patients (46.05%) showed allergic reaction to fruits, 64 (9.36%) of them showed a hypersensitive reaction to different kinds of fishes, and 528 patients (77.19%) had positive SPT results for different vegetables considered in our study, which included brinjal, cabbage, cauliflower, ladyfinger, spinach, and tomato.

In our present study, among the allergic population, maximum numbers of patients were hypersensitive to prawn (53.57%) and brinjal (45.67%) (Table 3). Brinjal was found to be most allergenic to patients belonging to the age groups 5 to 15 years (45.16%) and 41 to 60 years (55%), followed by prawn (5–15 years, 44.4%; 41–60 years, 50%). However, patients belonging to the age group 16 to 40 years showed maximum hypersensitivity to prawn (61.5%), followed by brinjal (43.42%). The allergenic reaction to prawn in this age group may be caused by greater consumption of prawn among individuals of this group. Similar results have been reported in patients in Russia, Estonia, and Lithuania.4 The percentage of allergy to beef was found to be quite high, although it is restricted only among the Muslim and Christian communities of Calcutta because beef consumption is not prevalent among the rest of the population. Thus, we can conclude that apart from age, food hypersensitivity also reflects different genetic factors and variations in cultural and dietary habits of each individual.19

ACKNOWLEDGMENTS

Sincere thanks to the University Grants Commission, India, for providing financial assistance to Jyotshna Mandal. The authors wish to thank Mr Ranjit Debnath and Mrs Rita Sen Sharma from the Institute of Child Health for their help in carrying out the SPT.

REFERENCES

1. Kumar R, Srivastava P, Kumari D, Fakhr H, Sridhara S, Arora N, et al. Rice (Oryzza sativa) allergy in rhinitis and asthma patients: A clinico-immunological study. Immunobiology. 2007;212:141–147.
2. Crespo JF, Rodriguez J. Food allergy in adulthood. Allergy; 2003;58:98–113.
3. Sampson HA. Update on food allergies. J Allergy Clin Immunol. 2004;113:805–819.
4. Eriksson NE, Moller C, Werener S, Magnusson J, Bengtsson U, Zolubas M. Self reported Hypersensitivity in Sweden, Denmark, Lithuania and Russia. J Investig Allergol Clin Immunol. 2004;14:70–79.
5. Hegde VL, Venkatesh YP. Anaphylaxis following ingestion of mango fruit. J Investig Allergol Clin Immunol. 2007;17:341–344.
6. Valenta R, Kraft D. Type 1 allergic reactions to plant derived food: a consequence of primary sensitization to pollen allergens. J Allergy Clin Immunol. 1996;97:893–895.
7. Mittag D, Vieths S, Vogel L, et al. Birch pollen-related food allergy to legumes: identification and characterization of the Bet v 1 homologue in mungbean (Vigna radiata), Vig r 1. Clin Exp Allergy. 2005;35:1049–1055.
8. Sharma S, Kathuria PC, Gupta KC, Nordling C, Ghosh B, Singh AB. Total serum immunoglobulin E levels in a case-control study in asthmatic allergic patients, their family members and healthy subjects from India. Clin Exp Allergy. 2006;36:1019–1027.
9. Patil SP, Niphadkar PV, Bapat MM. Chickpea: a major food allergen in the Indian subcontinent and its clinical and immunological correlation. Anu Allergy Asthma Immunol. 2001;87:140–145.
10. Parihar H, Kumar L, Puri Kumar V. The incidence of allergic diseases and feeding patterns in children up to 2 years of age. Indian J Pediatr. 1984;51:7–12.
11. Sharma J, Kumar L, Singh S. Allergenicity of common foods restricted in respiratory allergy. Indian J Pediatr. 2000;67:713–720.
12. Kumari D, Kumar R, Sridhara S, Arora N, Gaur SN, Singh BP. Sensitization to blackgram in patients with bronchial asthma and rhinitis: clinical evaluation and characterization of allergens. Allergy. 2006;61:104–110.
13. Sheldon JM, Lovell RG, Mathews KD, editors. A manual of clinical allergy. Philadelphia, PA: W.B. Saunders Co., 1967.
14. Dreborg S, Frew AJ, Eds: Allergen standardization and skin tests. WAO Journal • January 2009
15. Stytis DP, Stobo JD, Fudenberg H, Wells JV. Basic and clinical immunology. 4th ed. Singapore: Lange Medical Publishers, Maruzen Asia Pvt Ltd, 1992.
16. Bock SA, Atkins FM. Patterns of food hypersensitivity during sixteen years of double-blind, placebo-controlled food challenges. J Pediatr. 1990;117:561–567.
17. Kajosaari M. Food allergy in Finnish children aged 1-6 years. Paediatr Scand. 1984;51:713–720.
18. Kumari D, Kumar R, Sridhara S, Arora N, Gaur SN, Singh BP. Sensitization to blackgram in patients with bronchial asthma and rhinitis: clinical evaluation and characterization of allergens. Allergy. 2006;61:104–110.
19. Sampson HA. Relationship between food-specific IgE concentrations and the risk of positive food challenges in children and adolescents. J Allergy Clin Immunol. 1997;100:444–451.

TABLE 3. Allergens and the Total Number of Patients Positive to These Allergens

| Allergens | Total No. Patients Tested | No. Patients Positive (%) |
|-----------|---------------------------|---------------------------|
| Wheat (cereal) | 548 | 156 (28.47) |
| Egg | 498 | 174 (34.94) |
| Fish | 1094 | 600 (55.26) |
| Hilsa | 46 | 6 (4.55) |
| Prawn | 56 | 30 (53.57) |
| Rohu | 304 | 18 (11.84) |
| Fruits | | |
| Apple | 136 | 11 (8.09) |
| Banana | 330 | 134 (40.61) |
| Cucumber | 182 | 28 (15.38) |
| Guava | 123 | 31 (25.20) |
| Papaya | 236 | 62 (26.27) |
| Pumpkin | 254 | 49 (19.29) |
| Meat | | |
| Beef | 69 | 18 (26.08) |
| Chicken | 216 | 40 (18.52) |
| Mutton | 144 | 16 (11.11) |
| Milk | 554 | 114 (20.58) |
| Pulses | | |
| Musoor | 302 | 48 (15.89) |
| Mung | 126 | 24 (19.05) |
| Vegetables | | |
| Brinjal | 254 | 116 (45.67) |
| Cabbage | 214 | 36 (16.82) |
| Cauliflower | 284 | 82 (28.87) |
| Ladyfinger | 394 | 138 (35.02) |
| Spinach | 180 | 48 (26.67) |
| Tomato | 476 | 108 (22.69) |

© 2009 World Allergy Organization