Factors determining parents' perception of their child's risk of life-threatening food-induced anaphylaxis

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Factors that determine parents’ perception of their child’s risk of life-threatening food-induced anaphylaxis

Jennifer Ogg, M.B.B.Ch.,1 Jayne Wong, M.B.B.Ch.,1 Ming Wai Wan, Ph.D.,2 Naomi Davis, M.D.,3 and Peter D. Arkwright, M.D., Ph.D.1

ABSTRACT

Background: Although food allergy is known to be associated with increased disease burden, factors that shape parents’ perception of their child’s risk of future severe or fatal anaphylaxis are poorly understood.

Objective: This study aimed to evaluate factors associated with parents’ perceived risk of food-induced anaphylaxis.

Methods: A questionnaire-based survey of 202 parents was conducted in a single specialist center outpatient clinic that treats children with food allergies. Parents’ perceived risk of their child experiencing further food-induced anaphylaxis was assessed by using a validated food allergy independent measure. Demographic data as well as parents’ anxiety and depression scores were assessed by using the Hospital Anxiety and Depression score.

Results: Nineteen percent of parents believed that their child had a moderate-to-high chance of dying from food-induced anaphylaxis. A lack of a university education, higher anxiety score, and, particularly, possession of an epinephrine autoinjector (relative risk 9.5 [95% confidence interval, 1.7–53]) were key factors associated with heightened risk perception. Caring for a child with multiple food allergies was the main factor associated with parents feeling less able to manage future reactions (relative risk 9.9 [95% confidence interval, 3.3–30]). Parents’ risk perception of fatal anaphylaxis correlated with anxiety and mood scores.

Conclusion: Parents’ education, affect, and possession of an epinephrine autoinjector were associated with a heightened perceived risk of future anaphylaxis. Clinicians should consider not only the child’s needs but should also provide counseling for parents, particularly those who possess autoinjectors. Parents of children with multiple food allergies may need additional education and training to help them cope with future reactions.

Food allergy is common, occurring in ~8% of children in Western societies. Although there has been a sixfold increase in hospital admissions for food-induced anaphylaxis over the past 2 decades,2,3 deaths from anaphylaxis remain rare.4,5 Two concerns of both patients and their caregivers are the child’s risk of future anaphylaxis and their ability to effectively manage future reactions. Anxiety has a negative impact on quality of life (QoL) of both the patient and his or her family. Over the years, many guidelines have been published with the aim of reducing anxiety and engendering patient and caregiver confidence by providing information and advice on the management of allergies.5–16

Addressing the key factors associated with heightened levels of a patient’s and caregivers’ anxiety would reduce the overall disease burden.17,18 Although a number of previous studies sought to determine the nature of these factors, conclusions have been quite diverse. For example, a European study of 244 children with food allergy found that allergen type and nationality but not history of previous reactions or possession of an epinephrine autoinjector (EAI) were important in determining health-related QoL.19 A U.S. survey of 305 caregivers concluded that a Food Allergy Quality of Life–Parental Burden index was affected by not only allergen type but also the number of allergens, patient’s age, parental income, and history of reactions.20 A follow-up study of 3541 caregivers by the same U.S. group administered through social media reinforced these findings and indicated that QoL was better in parents of children who had received epinephrine for previous anaphylactic reactions, possibly by increasing the confidence of families that future reactions would be managed successfully.21 Cummings et al.22 indicated that there was reduced anxiety in children and parents who possessed an EAI, but their study was small and only included five parents who had no EAI. In con-

From the Institute of Inflammation and Repair, University of Manchester, Manchester, United Kingdom, Institute of Brain, Behaviour and Mental Health, University of Manchester, Manchester, United Kingdom, and Department of Paediatric Orthopaedic Surgery, Royal Manchester Children’s Hospital, Manchester, United Kingdom.

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Address correspondence to Peter D. Arkwright, M.D., Department of Paediatric Allergy and Immunology, University of Manchester, Royal Manchester Children’s Hospital, Oxford Rd., Manchester, M13 9WL, United Kingdom.

E-mail address: peter.arkwright@uhns.net

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An Australian survey found that QoL of 167 caregivers assessed by using the Food-Allergy Quality of Life scale was poorer in those caregivers who possessed an EAI. A subgroup of parents of children with food allergies, therefore, may benefit from additional support from health care professionals.

The aim of this study was to dissect out key psychological and physical factors associated with heightened parental perception of future anaphylaxis. As well as measuring levels of anxiety and depression, obsessive-compulsive tendencies of the parents were also assessed because we had previously observed a propensity to this personality trait in families with severe food allergies and atopic dermatitis. We hypothesized that a combination of demographic and psychological factors would be associated with parents’ risk perception. Primeau et al. found that caring for a child with peanut allergy was a greater disruption to family life than caring for a child with other diseases, e.g., arthritis. In this study, we compared the relative impact of a child with food allergy on parents’ future risk perception with a group of children who had sustained accidental bony injuries and who were attending an adjoining fracture clinic. Accidental injuries, although not usually chronic, are similar to food-induced anaphylaxis in that they are unpredictable and a cause of concern for most parents. Inclusion of this group allowed for a direct comparison of parents’ perceived risk of serious

### Table 1 Comparative questions in the questionnaire handed out to parents of children with food allergies and parents of children with injuries

<table>
<thead>
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<th>FAIM Questionnaire</th>
<th>Accidental Injury Questionnaire</th>
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<td><strong>Parents’ Perceived Risk</strong></td>
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<td><strong>The following questions are about the chance that you think something may happen to your child because of his or her food allergy; choose one of the answers provided; answer the questions by putting an “×” in the box next to the appropriate answer</strong></td>
<td><strong>The following questions are about the chance that you think something may happen to your child because of another accidental injury; choose one of the answers provided; answer the questions by putting an “×” in the box next to the appropriate answer</strong></td>
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<tr>
<td><strong>Questions</strong></td>
<td><strong>Questions</strong></td>
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<tr>
<td><strong>Chance that your child will accidentally eat something to which he or she is allergic?</strong></td>
<td><strong>Chance that your child will have another injury?</strong></td>
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<td><strong>Chance that your child will have a severe reaction if he or she eats something to which he or she is allergic?</strong></td>
<td><strong>Chance that your child will have a severe injury that needs a plaster cast or operation?</strong></td>
</tr>
<tr>
<td><strong>Chance that your child will die if he or she accidentally eats something to which he or she is allergic?</strong></td>
<td><strong>Chance that your child will die because of a future injury?</strong></td>
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<tr>
<td><strong>Chance that you will not be able to effectively deal with an allergic reaction should your child accidentally eat something to which he or she is allergic?</strong></td>
<td><strong>Response:</strong> Never (0% chance), very small chance, small chance, fair chance, great chance, very great chance, always (100% chance)</td>
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<td><strong>Impact on Child’s Life</strong></td>
<td><strong>Questions</strong></td>
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<td><strong>How great is the impact of your child’s food allergy on his or her school life?</strong></td>
<td><strong>How great is the impact of your child’s injury on his or her school life?</strong></td>
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<td><strong>How great is the impact of your child’s food allergy on his or her activities outside school?</strong></td>
<td><strong>How great is the impact of your child’s injury on his or her activities outside of school?</strong></td>
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<td><strong>Response:</strong> Almost none, very small, small, moderate, great, very great, extremely great</td>
<td><strong>FAIM</strong> = Food Allergy Independent Measure. Differences in wording between the two questionnaires are highlighted.</td>
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adverse events or death of their child with either food allergies, which are rarely fatal, and accidental injuries, the most common cause of death in school-age children.

METHODS

Study Design

This was a prospective, questionnaire-based study. The study had local ethics committee approval from the National Research Ethics Service, Liverpool Central Committee; all the parents provided informed written consent. Data sets were complete for all the participants, and no parents withdrew from the study. J. Ogg and J. Wong contributed equally to the study.

Patient Recruitment

Participants in the allergy group were recruited from parents of consecutive children who attended a specialist pediatric allergy outpatient clinic for assessment and management of food allergies. Clinicians at our specialist pediatric allergy service and throughout the northwest of England prescribe EAI if patients have a clinical history of anaphylaxis (respiratory or circulatory symptoms) to a food in which future avoidance is likely to be difficult (particularly peanut and tree nuts) or asthma is not controlled with low-dose inhaled steroids. Participants in the injury group were recruited from parents of children who attended an adjoining specialist pediatric fracture clinic for assessment and management of fractures and other bony injuries. Children who attended the fracture clinic and who had a history of food allergy, atopic dermatitis, asthma, or hay fever, or of chronic diseases that predisposed them to fractures were excluded from the study.

Data Collection

Parents were invited to complete a four-page standardized questionnaire. The questionnaire covered family demographics and details of the child’s food allergy or injury. Parents’ education and vocation were also recorded. Parental perceptions of the risk of future allergic reactions as well as information about disruption to daily life were collected by using the validated Food Allergy Independent Measure. A comparable questionnaire was used for parents with children who sustained injuries (Table 1). Anxiety and depression scores were measured by using the Hospital Anxiety and Depression Scale. Obsessive-compulsive tendencies were measured by using the Obsessive-Compulsive Inventory–Revised.

Statistical Analysis

Data from questionnaires were collated and analyzed by using IBM SPSS Statistics. Continuous vari-
and 48 (24%) had previously been admitted to the hospital because of their asthma.

**Parents’ Perceived Risk of their Child Experiencing Future Allergic Reactions**

Eighty-five percent of the questionnaires were completed by the mothers, 15% by the fathers. Demographic and psychological factors were not associated with the parents’ perceived future risk of their child eating food to which they were allergic (Table 2). However, parents of children with a history of an anaphylactic reaction and those who possessed an EAI were 2.6 times more likely to consider their child at a high risk of a future “serious and/or anaphylactic” reaction. Three times as many parents who possessed an EAI (12/70 [18%]) compared with those who did not have an EAI (7/132 [5%]) believed that their child was at high risk of a future anaphylaxis (Fig. 2). Possessing an EAI was the cofactor associated with the highest perceived risk of death from future allergic reactions (9.9 [95% confidence interval CI], 3.3–30; p < 0.001) (Table 2). One percent of the parents without an EAI (2/131) compared with 10% of the parents with an EAI (7/70) believed that their child was at high risk of death from a future allergic reaction (Fig. 2). The parents with higher anxiety scores (1.2 [1.1–1.4]; p = 0.02) and parents without a university education (3.7 [1.4–10.1]; p < 0.01) perceived that their child was at a higher risk of death from a future allergic reaction (Fig. 2). Nut allergy was not perceived by their parents as more likely to be associated with a high risk of serious or fatal anaphylaxis than allergies to other foods. Allergies to multiple foods was the only factor associated with parents’ perception that they would not be able to manage

<table>
<thead>
<tr>
<th>Cofactors</th>
<th>High Perceived Chance by Parents of</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Eating a Food To Which the Child Is Allergic</td>
<td>A Serious Future Allergic Reaction</td>
</tr>
<tr>
<td>RR (95% CI)</td>
<td>p Value</td>
<td>RR (95% CI)</td>
</tr>
<tr>
<td>Mother vs father</td>
<td>3.7 (1.0–13.9)</td>
<td>0.06</td>
</tr>
<tr>
<td>White European vs other ethnicity</td>
<td>1.3 (0.6–2.6)</td>
<td>0.5</td>
</tr>
<tr>
<td>University education</td>
<td>1.2 (0.6–2.3)</td>
<td>0.7</td>
</tr>
<tr>
<td>Age of child</td>
<td>1.0 (0.9–1.1)</td>
<td>0.6</td>
</tr>
<tr>
<td>Son vs daughter</td>
<td>0.6 (0.3–1.3)</td>
<td>0.2</td>
</tr>
<tr>
<td>Nuts vs other allergen</td>
<td>0.5 (0.2–1.1)</td>
<td>0.1</td>
</tr>
<tr>
<td>Multiple allergen groups</td>
<td>0.9 (0.4–1.8)</td>
<td>0.7</td>
</tr>
<tr>
<td>Anaphylaxis vs milder reaction</td>
<td>1.4 (0.6–3.1)</td>
<td>0.4</td>
</tr>
<tr>
<td>History of asthma</td>
<td>1.0 (0.5–2.2)</td>
<td>0.9</td>
</tr>
<tr>
<td>Possession of an autoinjector</td>
<td>0.8 (0.4–1.8)</td>
<td>0.6</td>
</tr>
<tr>
<td>Autoinjector used</td>
<td>3.1 (0.6–16.3)</td>
<td>0.2</td>
</tr>
<tr>
<td>HADS depression score</td>
<td>1.1 (0.9–1.2)</td>
<td>0.4</td>
</tr>
<tr>
<td>HADS anxiety score</td>
<td>1.1 (0.9–1.2)</td>
<td>0.3</td>
</tr>
<tr>
<td>OCIR score</td>
<td>1.0 (0.9–1.0)</td>
<td>0.3</td>
</tr>
</tbody>
</table>

RR = Relative risk; CI = confidence interval; HADS = Hospital Anxiety and Depression Scale; OCIR = Obsessive-Compulsive Inventory–Revised.

*High risk (great, very great, or 100% chance of a future event) compared with lower risk (0%, very small, small, or fair chance of a future event).

#Binary logistic covariant regression analysis with the addition of the variables.
future reactions (relative risk [RR] 7.1 [95% CI, 1.6–32]) (Table 2).

Because EAI possession was the main variable associated with a perceived risk of death from future anaphylaxis, further analyses that compared families with and those without EAI were performed (Table 3). Children who possessed EAI were significantly older (RR 1.1 [95% CI, 1.0–1.2]); were more likely to have had an anaphylactic reaction (RR 2.2 [95% CI, 1.1–4.6]), particularly to peanuts or tree nuts (RR 4.2 [95% CI, 1.9–9.1]); and to have allergic rhinitis. There were no significant difference in parents’ ethnicity, university education, or affect between the groups.

Psychological Factors and Parents’ Perceived Risk of Future Allergic Reactions in Children with Food Allergies

The mothers (n = 249) did not differ significantly from the fathers (n = 44) in any of the three psychology scores. In addition, these scores were not significantly different in parents of different educational attainment (data not shown). However, the parents who perceived that their child was at a higher risk of future allergic reaction to foods were significantly more anxious (Fig. 3). Those parents who believed that there was a risk of anaphylaxis or death were also more likely to have significantly higher depression and obsessive-compulsive disease scores.

Clinical Features of the Injury Group and Comparison with the Food Allergy Group

Of the 91 children who attended an adjoining pediatric orthopedic clinic for accidental sprains and fractures, 73 (80%) had sustained a fracture, 45% to a bone in their upper limb, 24% to their lower limb, and 2% to their pelvis or rib. Twenty-seven (30%) required an operation for the fracture or dislocation. Thirty-four (37%) had a history of an injury. The characteristics of children and their parents with food allergies and children and their parents with injuries are compared in Table 4. Children who presented with food allergies were significantly younger than those who presented with injuries (median age, 5 versus 10 years; p < 0.001) and were more likely to be white European (67% versus 52%; p < 0.01). Operative treatment was the only factor associated with a significantly higher perceived risk by the parents of “any” (3.9[1.4–11.0]; p = 0.01) or “serious” future injury (7.6 [1.3–44]; p = 0.02). Only 1 of 91 parents (1%) thought that there was a high chance of his or her child dying of a future injury. Parents’ perceived impact of their child’s allergy or injury on school life was similar, but parents whose children had had injuries believed that the injury had a significantly greater impact on the child’s home and extracurricular activities (Fig. 4). In contrast, parents of children with food allergies were more likely to consider their children at risk of future serious or fatal events (Fig. 5). The perceived risk of a serious future adverse event was 13-fold higher in the group with food allergies than in...
those with injuries \((p < 0.001)\), independent of the age of the child and the ethnicity of the parent.

**DISCUSSION**

To our knowledge, this is the first study to quantify parents’ risk perception of their child with serious or fatal allergic reactions to food. Half of the parents in this cohort believed that there was a moderate-to-high risk of their child having future anaphylaxis, and a fifth believed that their child had a moderate-to-high chance of dying from anaphylaxis. Previous anaphylaxis and possession of an EAI were two factors independently associated with a higher perceived risk of future anaphylaxis, after adjusting for confounders, such as the child’s age, type of food allergen, parents’ affect, and education, by using multivariate analysis. Peanut and/or tree nut allergy and a history of asthma were not associated with a higher perceived risk, even though there is published evidence for a link.\(^{31,32}\) The observed differences between the outcomes of our study and previous studies may relate to the measures used. We assessed parental risk perception, whereas others examined parental burden and/or QoL.\(^{17,19}\)

King et al.,\(^{33}\) examined the impact of peanut allergy on QoL and anxiety in 46 families but used SCAS or STAI rather than the Hospital Anxiety and Depression Scale to assess anxiety and found that the mothers had worse QoL and anxiety scores than the fathers. Although we found no significant difference in anxiety scores of the mothers and fathers, the \(p\) value was 0.055, and a larger sample number may have shown a significant trend. Another study, which compared parents in the Netherlands and the United States, showed that, although American parents were more likely to visit a physician for their child’s food allergies and had a better knowledge of food avoidance strategies than the Dutch parents, the U.S. parents had lower QoL scores.\(^{34,35}\)

Parents who thought that their child was at a high risk of dying from anaphylaxis were less likely to be university educated. They also tended to have higher anxiety scores. A possible explanation for the latter observation is that having a child with food allergies makes parents more anxious. Gillespie et al.\(^ {36}\) in an interview-based study, indicated that consultation with an allergy specialist may indeed initiate or heighten maternal anxiety about the impact of their child’s food allergies. Alternatively, more anxious parents might worry more about the risks of food allergies, but, if this were the case, then one might expect that they might also request and receive EAI’s from their child’s physician. Our study found that parents in possession of an EAI were not more anxious than those without an autoinjector.

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>No EAI ((n = 132))</th>
<th>EAI ((n = 70))</th>
<th>(p) Value*</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age, median (IQR), y</td>
<td>4 (2–8)</td>
<td>8 (4–11)</td>
<td>0.001</td>
</tr>
<tr>
<td>Boys, no. (%)</td>
<td>76 (58)</td>
<td>45 (64)</td>
<td>0.3</td>
</tr>
<tr>
<td>An only child, no. (%)</td>
<td>76 (58)</td>
<td>45 (64)</td>
<td>0.4</td>
</tr>
<tr>
<td>White European origin, no. (%)</td>
<td>89 (67)</td>
<td>47 (67)</td>
<td>1.0</td>
</tr>
<tr>
<td>Anaphylactic symptoms, no. (%)</td>
<td>26 (20)</td>
<td>31 (44)</td>
<td>0.001</td>
</tr>
<tr>
<td>EAI previous used, no. (%)</td>
<td>0 (0)</td>
<td>8 (11)</td>
<td>0.001</td>
</tr>
<tr>
<td>Peanut or tree nut allergy, no. (%)</td>
<td>51 (39)</td>
<td>55 (79)</td>
<td>0.001</td>
</tr>
<tr>
<td>Cow’s milk protein allergy, no. (%)</td>
<td>53 (24)</td>
<td>23 (33)</td>
<td>0.2</td>
</tr>
<tr>
<td>Atopic diseases, no. (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Atopic dermatitis</td>
<td>110 (83)</td>
<td>48 (69)</td>
<td>0.02</td>
</tr>
<tr>
<td>Asthma</td>
<td>46 (35)</td>
<td>34 (49)</td>
<td>0.07</td>
</tr>
<tr>
<td>Allergic rhinitis</td>
<td>32 (25)</td>
<td>30 (51)</td>
<td>0.001</td>
</tr>
<tr>
<td>Questionnaire completed by the mother, no. (%)</td>
<td>109 (83)</td>
<td>66 (94)</td>
<td>0.03</td>
</tr>
<tr>
<td>Parents with university education, no. (%)</td>
<td>65 (49)</td>
<td>27 (39)</td>
<td>0.4</td>
</tr>
<tr>
<td>Parents’ psychological profile, median (IQR)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>HADS depression score</td>
<td>4 (1–7)</td>
<td>3 (1–6)</td>
<td>0.7</td>
</tr>
<tr>
<td>HADS anxiety score</td>
<td>5 (2–9)</td>
<td>6 (4–8)</td>
<td>0.3</td>
</tr>
<tr>
<td>OCD score</td>
<td>7 (3–15)</td>
<td>6 (4–12)</td>
<td>0.6</td>
</tr>
</tbody>
</table>

*Compared patients who had and patients who had not been prescribed an EAI.

#Anaphylaxis is defined as allergic reactions associated with respiratory or circulatory symptoms.

**Statistical analysis** was performed by using either the \(\chi^2\) test for discrete variables or the Mann-Whitney \(U\) test for continuous variables.
EAsIs are prescribed to treat anaphylaxis that could potentially, albeit rarely, lead to death. Our criteria for EAI prescribing, detailed in the Methods section, are more limited than those indicated in the 2015 U.S. anaphylaxis practice parameter update. These practice parameters not only recommend EAIs for all patients who have experienced or are at risk for food-induced anaphylaxis (including patients with food allergy and asthma; patients allergic to peanut, tree nuts, fish, and shellfish; and patients with a history of a systemic reaction to food) but also indicated that consideration be given to prescribing EAIs to all patients with immunoglobulin E–mediated food allergy. Our study indicated that parents who possessed EAI not only perceived that their child was at risk of future anaphylactic reactions but of fatal reactions. They supported those of Pinczower et al. who, in a recent Australia survey of 103 parents of children with food allergies, found that EAI possession was associated with significantly worse food allergy-related QoL scores and contrasted to other reports that indicated that parents with EAI are more confident in managing allergic reactions. Poor confidence may relate to lack of hands-on experience and fear of using EAIs. Further studies are required to directly compare EAI competence and confidence in parents who had and parents who had not used an EAI to treat anaphylaxis.

Compared with the injury group, parents of children with food allergies were 13-fold more likely to consider that their child at a high risk of a future serious adverse event and 21-fold more likely to consider that their child may die because of a future adverse event, independent of the child’s age, sex, and ethnicity. This was at odds with pediatric morbidity and mortality statistics, in which accidental injuries account for one-third of all emergency department visits and are the most common cause of death in school-age children and young people worldwide, whereas allergic reactions to food accounts for <1% of visits to emergency departments and only one to two childhood deaths in the United Kingdom each year. The reasons for using a group of children with bony injuries for comparison with children with food allergies deserves further clarification. First, both food-induced anaphylaxis and injuries are accidental. Second, as detailed above, accidental injuries are also potentially life threatening. Third, although not usually leading to chronic disease, injuries are potentially recurrent. The modified Food

Figure 3. The association between the parents’ affect and obsessive-compulsive disease (OCD) scores and the perceived risk that their child with a food allergy will have a future allergic reaction. The Hospital Anxiety and Depression Scale anxiety and depression scores as well as obsessive-compulsive scores of parents in relation to their perception that the child was at future risk of death (top panels), of a serious adverse event (middle panels), or of eating something to which he or she is allergic (lower panels). Data are shown as box plots (median and interquartile ranges) for low risk, moderate-to-fair risk, and high risk bars. Comparisons were made between median scores of parents who perceived a low and moderate or a high risk of future events for their children by using Kruskal-Wallis tests. *p < 0.05, **p < 0.005 (2-tailed test).
Allergy Independent Measure questionnaire for perception of the future risk of children with bony injuries has not been previously validated. However, of the observed trends in perceived risk of “any injury” (28%), a “serious injury” (9%), and “death” (1%), the higher perceived impact of injuries on life outside of school compared with within school and the higher perceived risk in children who require an operation, all provided evidence for the validity of the questions.

**CONCLUSION**

It is understandable that some parents are unsure, concerned, or even overwhelmed by their child’s food
Discuss allergies. This study highlighted the discrepancy between perceived and actual risks that some parents have about their child with food allergies. Parents who are more anxious or have not been to university may need additional support to focus on the evidence, particularly with regard to the actual risk of dying from anaphylaxis and how to manage children with multiple allergies. A prescription of EAI should not only be accompanied by an explanation as to when and how to use the device but a clear explanation of the actual risks that physicians hope to avert by prescribing the device.

ACKNOWLEDGMENTS

The authors thank Timothy David, University of Manchester, Richard Pumphrey, honorary clinical immunologist, Manchester Royal Infirmary, and Allie Crewe, parent representative, for their helpful comments regarding this manuscript.

REFERENCES


AUTHOR QUERIES

AUTHOR PLEASE ANSWER ALL QUERIES

1—Au: Please check all author names for accuracy. For indexing purposes, please indicate the surname (family name) of each author. In addition, please confirm that authors are listed in the correct order and that all names are spelled correctly.

2—Au: Please verify. Okay to add “95% confidence interval” for “1.7-53”

3—Au: In Table 1: Please note. Journal style is to not use highlighting. Please use bold and rephrase “Differences in wording between the two questionnaires are highlighted.”

4—Au: Please add company information, including complete company name, city, and state (or country), for “IBM SPSS Statistics”

5—Au: In Table 2: Please clarify “Eating a food to which the child is allergic” is this “Of the child eating a food to which her or she is allergic” ? “A serious future allergic reaction” - is this “The child having a serious future allergic reaction” ? “Death from future allergic reaction” - is this “The child’s death from a future allergic reaction” ? “Able to manage future allergic reaction” - is this “Of being able to manage the child’s future allergic reaction” ? Please indicate what the data in bold represent. Please verify. Are the footnote designators okay with the table title? Or, move to appropriate places within the table. Please note. Journal style indicates that p cannot equal 0. Okay as is?

6—Au: Please clarify. Is this “RR 9.9” ?

7—Au: Please verify. Is this “RR” and “95% CI” - “1.2 [1.1-1.4]”

8—Au: Please verify. Is this “RR” and “95% CI” - “3.7 [1.4-10.1]”

9—Au: Please move “relative risk” to first mention in the text if necessary.

10—Au: In Table 3: Please indicate what the data in bold represent. Please note. AMA Journal style indicates that p cannot equal 1.0. Okay as is? Please verify. Please indicate where footnotes # and § apply within the table.

11—Au: In Table 4: Please indicate what the data in bold represent. Please verify. Is p = 1.0 okay as is ? Please indicate where the footnotes # and § apply within the table.

12—Au: Please verify. Is this “RR: and “95% CI” - “(1.4-11.0)” and “(1.3-44)”?

13—Au: Please verify. Okay to add “To our knowledge,” ?

14—Au: Please spell out “SCAS” and “STAI”

15—Au: Please clarify. Is this “EAI” - “an autoinjector” ?
AUTHOR QUERIES

AUTHOR PLEASE ANSWER ALL QUERIES

16—Au: Please verify. Okay as spelled out - “IgE”?

17—Au: Please identify “They”

18—Au: Please note. This reference was not located in PubMed. Please verify/correct all information.

19—Au: Please clarify “Low”

20—Au: Please add a complete citation for “Zoa et al, 2002”

21—Au: Please add a complete citation for “Zoa et al, 2002”

22—Au: Please add department(s) and/or division(s) for affiliations 1 and 2.

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