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13 **A cross-sectional, population-based study on the prevalence of food allergies among**
14 **children in two different socio-economic regions of Vietnam**

15

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30

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41

42 **Number of tables and figures**

43 **Table 1.** Demographics of participating children in Hue and Tien Giang

44 **Table 2.** Comparison of the prevalence of self-reported FA and doctor-diagnosed FA in two
45 survey populations

46 **Table 3.** Multivariable logistic regression analysis of demographic factors for FA

47 **Figure 1.** Proportion of reported clinical symptoms in participating children in Hue and Tien
48 Giang. **A.** Reported food-induced adverse symptoms ($n = 2,905$). **B.** Reported adverse
49 symptoms in self-reported FA participants ($n = 763$). **C.** Reported adverse symptoms in
50 doctor-diagnosed FA participants ($n = 580$).

51 **Figure 2.** Comparison of the distribution of reported food groups eliciting clinical reactions
52 in participating children in Hue and Tien Giang: **A.** Reported food-induced adverse
53 symptoms in Hue (number of participants $n = 911$); **B.** Self-reported FA in Hue; **C.** Doctor-
54 diagnosed FA in Hue; **D.** Reported food-induced adverse symptoms in Tien Giang; **E.** Self-
55 reported FA in Tien Giang; **F.** Doctor-diagnosed FA in Tien Giang. The total number of
56 reported food groups is presented for each study area and symptom group.

57

58 **Additional file**

59 **Appendix S1.** Survey Questionnaire for child participants

60

61 **Ethics approval**

62 This study was approved by the Human Research Ethics Committee (HREC) at James Cook
63 University (ID: H6437).

64

65 **Consent for publication**

66 All authors have approved the manuscript for submission.

67

68 **Availability of data and materials**

69 The datasets used and/or analysed during the current study are available from the
70 corresponding author on reasonable request.

71

72 **Competing interests**

73 The authors declare that they have no competing interests.

74

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79 Food and Allergy Research, Australia.

80

81 **Authors' contributions**

82 TL and AL developed the concept and study design. DN and TL conducted the on-site
83 survey. AV and TL processed survey data. TL performed the statistical analysis. TL wrote

84 the manuscript. AT, TR and AL edited the final manuscript. All authors contributed to the
85 development of the manuscript and approved the final version.

86

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88 **ABSTRACT PAGE**

89

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96

97 **A cross-sectional, population-based study on the prevalence of food allergies among**
98 **children in two different socio-economic regions of Vietnam**

99

100 **Pediatr Allergy Immunol**

101

102 **Abstract**

103 **Background:** There is a paucity of data on the prevalence of food allergy (FA) in Vietnam. A
104 cross-sectional, population-based study was conducted to evaluate the current prevalence of
105 FA among 2-6-year-old children in two different regions in Vietnam.

106 **Method:** A structured, anonymous questionnaire, modified from published FA
107 epidemiological studies and based on EAACI guidelines, was distributed to parents/guardians
108 of participating children in Hue City (urban area) and Tien Giang Province (rural area). Data
109 collected from the survey were statistically analyzed to generate the prevalence of self-
110 reported and doctor-diagnosed FA and overarching pattern of food allergens.

111 **Results:** A total of 8,620 responses were collected (response rate 81.5%). Children in Tien
112 Giang reported more than twice the food-induced adverse reactions seen in children in Hue
113 (47.8% vs. 20.5%). In contrast, children in Hue showed higher self-reported FA (9.8%) and
114 doctor-diagnosed FA rates (8.4%) than children in Tien Giang (7.9% and 5.0%, respectively).

115 Crustacean was the predominant allergy-inducing food in both areas (330 of 580 cases,
116 56.9%), followed by fish, mollusk, beef, milk and egg. However, substantial variations of FA
117 patterns were seen between the study sites. Geographic location and co-morbidities of other
118 allergic diseases were key risk factors of FA ($P<0.001$).

119 **Conclusions:** The prevalence of FA in Vietnamese children seems to be higher than
120 previously reported from other Asian countries. Crustacean is the predominant allergy-
121 inducing food among participating preschool children in Vietnam. The variation of reported
122 food allergen sources across different socio-economic locations could imply different eating
123 habits or the participation of indoor and outdoor allergen exposure.

124

125 **Key words:** children, crustacean allergy, fish allergy, food allergy, prevalence, Vietnam.

126

127 **Abbreviations**

128 FA: Food allergy

129 EAACI: European Academy of Allergy and Clinical Immunology

130 IgE: Immunoglobulin E

131 CI: Confidence Interval

132 OFC: Oral Food Challenge

133

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146

147 **Introduction**

148 Food allergy (FA), an adverse immune reaction to food proteins, has a wide spectrum of
149 clinical presentations, ranging from mild skin problems to severe systematic reactions. In the
150 most severe case, FA can lead to anaphylaxis and might result in death within minutes. FA is
151 estimated to affect about 8% of children and 5% of adults in the general population
152 worldwide (1).

153

154 Children are more likely to develop food allergies than adults due to remaining controversial
155 causes, including the immature immune system in childhood and/or the inappropriate food
156 introductory practices (2, 3). Eight food groups, often referred as the “Big 8”, account for
157 over 90% of food allergic reactions and include cow’s milk, egg, peanut, tree nuts, soy,
158 wheat, fish and shellfish. Except for cow’s milk and egg allergy which are often outgrown,
159 most other FAs often persist for life (1). So far, no cure is available and childhood FA
160 imposes a substantial health and economic burden for children and their caregivers.

161

162 The common food commodities accounting for FA in children are cow’s milk, egg, peanut,
163 tree nuts and fish (4), while the first three foods are the leading causes for paediatric
164 anaphylaxis in Western countries (5). In Asia, the prevalence of paediatric FA seems to vary
165 between 1.11-7.65% (6), and the patterns of FA showed marked difference from other parts
166 of the world (1). Recent studies among 2-7-year-old children from Singapore, Thailand, the
167 Philippines and Hong Kong demonstrated that shellfish allergy was dominant, but not milk,
168 egg or peanut (7, 8). Furthermore, fish was reported to be the predominant allergen in
169 adolescents in the Philippines, Singapore and Thailand (9). Within Asia, studies from Japan
170 and Korea showed different FA patterns to most available FA data, with wheat allergy
171 particularly common in East Asian countries (6). These data were supported by a recent study
172 from Australia, where large differences in allergy/anaphylaxis risk and trigger were
173 demonstrated between children born in Australia and children born in Asia (10). The

174 variation of FA patterns throughout Asia indicates that region specific and accurate data on
175 FA prevalence and clinical patterns are crucial for an effective FA management program in
176 any community.

177

178 In Vietnam, about 4.4 million children aged 2-6-year-old attend kindergartens, accounting
179 over 90% of all children at this age group in 2016 (11). The first population-based study on
180 FA in adults was recently performed by our group, revealing a high rate of FA in this
181 population: self-reported FA (18.0%) and doctor-diagnosed FA (5.8%) (12), while no data on
182 children is available so far. We sought to evaluate the epidemiologic and clinical features of
183 FA in Vietnamese preschool children. The possible variations of childhood FA prevalence
184 and its associated risk factors in socio-economically different regions in Vietnam were also
185 investigated.

186

187 **Methods**

188 **Study design and subjects**

189 A cross-sectional, population-based study was conducted in preschool children aged from 2
190 to 6 years in 2016. Survey participants were randomly selected using the cluster sampling
191 method from a list of 25 kindergartens in Hue city and 14 kindergartens in Cai Be district,
192 Tien Giang province, representing a total of 104,602 preschool children in two regions (11).
193 The paper-based questionnaires were distributed to parents/guardians of children at their
194 kindergartens. Most of the answer sheets were collected on the same day. The response rate
195 was calculated based on the number of returned answer-sheets divided by the total distributed
196 questionnaires.

197

198 **Sample size calculation**

199 To obtain a statistical estimation of the prevalence of FA, the minimum sample size was
200 calculated based on the current estimated prevalence of FA in children (8%) in the general
201 population (13); the chosen precision of the estimation $d=1/5p$ was calculated with a
202 statistical confidence of two standard errors of the mean $z=1.96$ (95% Confidence Interval
203 (CI), $P<0.05$). The minimum necessary sample size calculated for children was 1,825
204 participants.

205

206 **Study locations**

207 The study was conducted in two different regions of Vietnam: Hue City and Cai Be District
208 of Tien Giang Province. Hue City is in the Central region of Vietnam with a population
209 density of 5,011 per square kilometer. The main economic activities in Hue are tourism,
210 industry and aquaculture. Urbanisation has quickly taken place in this city due to the rapid
211 development of tourism. Hue has an average temperature of 25.4°C, average humidity of 87%
212 and a total of 1,754.2 hours of sunshine per year.

213 Cai Be District is a rural area in the Mekong Delta of southern Vietnam. This river-land
214 mixed town has a population density of 657 per square kilometer. The major economic
215 activities in Cai Be are aquaculture, rice and fruit farming. Cai Be-Tien Giang has an average
216 temperature of 28.2°C, average humidity of 80.4% and a total of 2,104.6 hours of sunshine
217 per year (14).

218

219 In this study, taking into consideration the effects of population density, living lifestyle and
220 environmental conditions, we defined participants in Hue City as living in urban area and
221 participants from Cai Be District as living in rural area.

222

223 **Questionnaire design**

224 The questionnaire, modified from published studies in the US and Asia (7, 9), had two parts:
225 part I asked the participant demographic information, and part II contained ten questions on
226 FA (Appendix S1, Supporting Information). The questionnaire was translated into
227 Vietnamese. The content of the questionnaire and its translation were reviewed and approved
228 by the Human Research Ethics Committee at James Cook University (ID: H6437). By
229 answering the questionnaire, the parents/guardians gave the informed consent to the study
230 and the permission to use obtained child health information for research publications and
231 reports.

232

233 **Definitions**

234 We established a set of criteria to define self-reported and doctor-diagnosed FA in this survey
235 based on the most recent European Academy of Allergy and Clinical Immunology (EAACI)
236 guidelines on FA and anaphylaxis (15). In specific, the suggestive symptoms of FA were
237 considered including persistent symptoms towards food ingestion and the co-occurrence of
238 two or more different clinical presentations (16). The typical symptoms for IgE-mediated FA
239 included hives/urticaria or angioedema or vomiting or gastrointestinal symptoms or
240 anaphylactic reactions (i.e. reduced blood pressure, loss of consciousness, chest pain and

241 weak pulse) after food intake. In this study, children with only one symptom of
242 hives/angioedema were also defined as food allergic and included.

243 *Self-reported FA* was the group of participants who fulfilled the above criteria and reported
244 having FA.

245 *Doctor-diagnosed FA* was the group of participants with self-reported FA, which was
246 clinically confirmed by a medical practitioner.

247 *Food-induced adverse symptoms*: any abnormal clinical response that occurs following
248 ingestion of a food or food component.

249 *Family history of FA* was defined when the participant had in their immediate family a
250 member with FA.

251 *Coexisting other allergic diseases* was defined when the participant had any other allergic
252 diseases including pollen allergy, antibiotic allergy, asthma, eczema etc.

253

254 **Statistical analysis**

255 The survey data were analyzed and plotted using the IBM SPSS Statistics for Windows,
256 version 24.0 and GraphPad Prism version 7.03. Continuous variables were presented as
257 median and interquartile range (IQR). Categorical data were compared by using either
258 Fisher's exact test or Chi-square test with a 2-tailed *P*-value. The Wilson/Brown method was
259 performed to provide a 95% CI of proportions. Multiple logistic regression model was used
260 to study the association between multiple risk factors and the incidence of having doctor-
261 diagnosed FA. A *P*-value of <0.05 was considered as statistically significant for all tests.

262 **Results**

263 **Participants**

264 A total of 8,620 questionnaires were completed and returned from the two survey sites
265 (response rate 81.5%). The survey in Hue gained a higher response rate (93.5%) than in Tien
266 Giang (69.5%). Minimal difference in gender distribution was observed across the two survey
267 sites. The age median (IQR) of the participants was 4 (2-6) years in Hue and 6 (2-6.5) years
268 in Tien Giang. The demographic characteristics of participating children are presented in
269 Table 1.

270

271 **Comparison of reported food-induced adverse symptoms between children in Hue and** 272 **Tien Giang**

273 Children in Tien Giang were reported to have twice the food-induced adverse symptoms than
274 children in Hue (47.8% vs. 20.5%) (Table 1). However, self-reported FA in Hue (9.8%) was
275 higher than in Tien Giang (7.9%) (Table 2). In the perceived FA group, more children in Hue
276 presented to doctors for medical advice, 91.6% compared to 76.7% in Tien Giang. Overall,
277 the prevalence of life-time doctor-diagnosed childhood FA in Hue was 8.4%, nearly double
278 the rate of 5.0% in Tien Giang ($P<0.0001$).

279
280 Suspected FA children in Hue reported less concurrent episodes than those in Tien Giang (an
281 average of 1.4 episodes compared to 2.0 episodes, respectively). Hives, diarrhea and nausea
282 or vomiting were the most predominant clinical presentations reported. Ten participants
283 (0.2%) in Tien Giang experienced severe symptoms (i.e. loss of consciousness, drop in blood
284 pressure, chest pain and weak pulse) due to FA while in Hue, only one case was reported
285 (0.02%) (Figure 1).

286

287 **Distribution of the major food allergens in FA children in Hue and Tien Giang**

288 Most of the affected subjects (78.1%) reported food adverse symptoms to only one food item;
289 13.1% reported adverse reactions to two different food items and 7.1% had reactions to more
290 than two different food groups. Crustacean was the most predominant allergy causing food
291 type in both Hue (50.1%) and Tien Giang (30.6%), while the distribution of the remaining
292 'Big 8' food groups were very different (Figure 2). Statistically significant differences were
293 seen in the prevalence of crustacean, mollusk, beef, milk, wheat and tree nut allergies
294 between children in Hue and Tien Giang ($P<0.05$) (Table 2).

295

296 **Contribution of environmental factors to FA incidence**

297 Genetic and environmental factors are reported to play a role in the development of FA (1,
298 17). In this study, we analyzed the contribution of geographical location, gender and family
299 history of FA as well as coexisting other allergic diseases to the FA incidence by using
300 multivariable logistic regression model. A strong influence of participant location and atopic
301 conditions to FA risk was observed in this study. Children living in Hue (urban area) have a
302 higher risk of having FA than children living in Tien Giang ($OR: 3.902, P<0.001$). The FA

303 rate was found to be 3.428 times higher in participants with other existing allergic diseases
304 ($P < 0.006$). Gender and family history of FA showed no impact on FA risk in this study
305 population (Table 3).

306

307 **Discussion**

308 This population-based survey is the first to establish the prevalence of self-reported FA
309 (8.9%) and doctor-diagnosed FA (6.7%) in Vietnamese children. Our findings indicate large
310 variations of FA prevalence between two survey sites with different socio-economic
311 backgrounds. The population living in the urban area presented a higher prevalence of FA but
312 also had a higher rate of doctor consultation to diagnose FA. Most participants (78.1%)
313 reported adverse symptoms to only one food group, with crustacean the dominating food
314 allergen. Hives and gastrointestinal tract problems were the most commonly reported clinical
315 symptoms for both regions.

316

317 We observed a higher rate of self-reported FA (8.9%) than doctor-diagnosed FA (6.7%),
318 consistent with previous assessments of questionnaire-based FA rates in Asian populations
319 (1.11 - 7.65%) (6). This variation appears to be determined by the complex pathophysiology
320 of adverse reactions to food and the perception of respondents of this disease. Common
321 aetiology in paediatrics with food-related adverse symptoms are immune-mediated FAs and
322 non-immune mediated food intolerance (18). There is lack of strong evidence to differentiate
323 FA from food intolerance exclusively based on reported clinical history, especially in Asian
324 communities. Among doctor-diagnosed milk allergic participants, two thirds of participants
325 presented gastrointestinal symptoms which might imply the contribution of other food-
326 induced disorders rather than true FA. A food outbreak, suspected to be an acute allergic
327 reaction to a new formula product, was recorded in 19 out of 229 hospitalised children in
328 2009 (19). Unfortunately, no allergens were identified due to the constraint of diagnostic
329 capacity in Vietnam. Further investigations will exclude other non-Immunoglobulin E (IgE)-
330 mediated FAs, such as Food Protein-Induced Enterocolitis Syndrome and eosinophilic
331 esophagitis in the paediatric population to give an accurate estimation of true FA prevalence
332 (20).

333

334 Patient's clinical history of FA is the initial motive for further diagnostic analysis, however,
335 only 4 to 5% of the self-reporting FA population is generally confirmed as true FA (4).
336 Parent-reported FA in Thai children was found to be 9.3%, but reduced to 1.1% when
337 confirmed by oral food challenge (OFC) (21). A survey of Singapore-born children aged 4-6
338 years showed the variation of self-reported FA to shellfish with 7.22 % as compared to a rate
339 of 1.19 % with convincing history FAs (7). As it was consistently concluded in previous
340 studies, an accurate diagnostic procedure of IgE-mediated FA must comprise of multiple tests
341 including skin prick testing, measurement of serum specific IgE and OFC (22). However,
342 only limited services are available in Vietnam for diagnosing FA, particularly in rural areas.
343 Most commercial diagnostic tests that are readily available in Western countries, including
344 IgE quantification and skin prick tests, are not registered or partially available to private
345 patients and in specialized clinics. In the presented study, data could not be collected for the
346 onset of adverse symptoms that might have better supported differentiating between IgE-
347 mediated and non-IgE mediated FA. This is one of the biggest challenges in studying the
348 prevalence of FA in a country where only a few people have access to correct FA diagnosis.
349 This paper-based survey on health conditions was thought to be a rather new practice for
350 most Vietnamese, so we aimed and succeeded at keeping the questionnaire as simple as
351 possible to achieve a high response rate (81.5%).

352

353 This study revealed a distinct distribution of the "Big 8" food allergens in Vietnamese
354 children. Unlike the patterns of childhood FA from Western populations, previous studies in
355 Asian populations showed the predominance of shellfish and fish allergy rather than egg,
356 cow's milk and peanut (6, 7, 9), and this tendency was also determined in this survey.
357 Children from rural and urban Vietnam reported higher adverse reaction rates to seafood,
358 then beef, milk and egg. The predominance of seafood allergy in Asia might be claimed for
359 the availability and high consumption of this food commodity (23). In Vietnam, the average
360 fish consumption is with 33 kg per capita per annum much higher than the world's average
361 consumption of 21 kg (23). The impact of ethnic characteristics to seafood allergy in Asian
362 communities was validated in a study among expatriate and local Singaporean children,
363 revealing the predominance of shellfish allergy in local children compared with expatriate
364 children (7).

365

366 Considering ethnic characteristics and cultural dietary practices, we found considerable
367 variations of FA prevalence among urban and rural population in Vietnam. Crustacean and
368 milk allergy are predominant in children in Hue (urban area). However, there was insufficient
369 data on the consumption of these commodities between the two areas to postulate FA risk.
370 The high incidence of shellfish allergy in urban children might be related to higher exposure
371 to indoor allergens as discussed in the current literature (24). For instance, indoor mites were
372 documented to cross-react with the major shellfish allergen tropomyosin (25), and storage
373 mites were identified in indoor environments in the north of Vietnam (26). In contrast,
374 children in the Tien Giang province showed a much higher prevalence to mollusk, wheat, tree
375 nuts and beef. Recent studies in the US and Sweden documented the association of red meat
376 allergy to tick bites (27, 28), which was explained by the cross-reactivity of a carbohydrate
377 oligosaccharide galactose- α -1,3-galactose in mammalian meat and a similar component
378 found in the saliva of tick. Children from rural areas are more likely to have tick bites than
379 those in the city (29), and therefore environmental factors might contribute to the high rate of
380 beef allergy in children in Tien Giang. Similarly, the high incidence of wheat and tree nut
381 allergy in this subpopulation might be explained by the possible cross-reactivity of these food
382 allergens with other aeroallergens abundant in the rural area. It should be noted that wheat is
383 not a staple food in Vietnam and no data on gluten intolerance or Coeliac disease have been
384 reported so far in this population. This will be of interest for further investigation on the
385 influence of environmental factors to FA.

386
387 The data from the multivariable logistic regression analysis of demographic risk factors
388 (gender, family history of FA, coexisting other allergic diseases and geographic location)
389 demonstrated a strong contribution of coexisting other allergic diseases ($OR= 3.428$,
390 $P<0.006$) to FA incidence, but not a family history of FA ($OR=1.018$, $P=0.961$). FA is
391 thought to run in a family (17). However, the contribution of a family history of FA to the
392 risk of FA development remains inconsistent among studies (30, 31). In the present study, we
393 did not apply any additional logistic regression models to further assess individual risk
394 factors for FA.

395
396 The strengths of this study are the large population-based dataset ($n = 8,620$) collected at two
397 different socio-economical survey sites and the high response rate (81.5%). The limitations of

398 this study are the self-administered data on FA and therefore the response might contain
399 recall bias. Our target population were children age from 2-6 years and the information on
400 children outside this age group with potentially different FA rates have not been included.
401 There are several factors such as the disparity of the medical facilities among rural and urban
402 areas in Vietnam and the economic circumstances of participants that might contribute to the
403 variation on reported FA rates among the two study sites.

404

405 In conclusion, this study contributes to the current paucity of FA data in the broader Asian
406 population and is the first to profile this emerging epidemic in Vietnam. Our study clearly
407 showed that FA is prominent in Vietnam, but unexpected patterns of food allergies are
408 perceived. A large variation of FA incidence was observed in subpopulations from rural and
409 urban regions, implying possible impacts of living conditions. Further investigations are
410 necessary to confirm the true prevalence of FA and possible cross-reactivities between
411 different allergen sources for a precise diagnosis and better management of this serious
412 childhood illness.

413

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496

497 **TABLE 1.** Demographics of participating children in Hue and Tien Giang.

Variable, <i>n</i> (%)	Hue	Tien Giang	Difference, <i>P</i>	Total study population
Total	4,443	4,177		8,620
Female	2,206 (49.6)	2,120 (50.8)	0.2860	4,326 (50.2)
Male	2,239 (50.4)	2,055 (49.2)	0.2860	4,294 (49.8)
Age group (years)				
2 to <3	1,140 (25.7)	52 (1.3)	<0.0001	1,192 (13.8)
3 to <4	1,365 (30.7)	655 (15.7)	<0.0001	2,020 (23.4)
4 to 6	1,940 (43.6)	3,467 (83.0)	<0.0001	5,407 (62.7)
Age, Median (IQR)	4 (2-6)	6 (2-6.5)		
Reported adverse reactions to food	911 (20.5)	1,994 (47.8)	<0.0001	2,905 (33.7)
Self-reported FA	433 (9.8)	330 (7.9)	0.0026	763 (8.9)
Seeking medical advice for FA [‡]	394 (91.6)	250 (76.7)	<0.0001	644 (84.4)
Doctor-diagnosed FA	373 (8.4)	207 (5.0)	<0.0001	580 (6.7)
<i>FA to 1 food group</i>	328 (87.9)	125 (60.4)	<0.0001	453 (78.1)
<i>FA to 2 different food groups</i>	40 (10.7)	36 (17.4)	0.9084	76 (13.1)
<i>FA to more than 2 different food groups</i>	4 (1.1)	37 (17.9)	<0.0001	41 (7.1)

498 [‡] Among subjects with self-reported FA.

499 The Fisher's exact test was performed using GraphPad Prism for Windows (GraphPad
500 Software, La Jolla California USA) to obtain *P* values.

	Self-reported FA				Doctor-diagnosed FA			
	Hue	Tien Giang	Difference, <i>P</i>	Entire study population	Hue	Tien Giang	Difference, <i>P</i>	Entire study population
Any food	9.75 (8.91 – 10.65)	7.90 (7.12 – 8.76)	0.0027	8.85 (8.27 – 9.47)	8.40 (7.62 – 9.25)	4.96 (4.34 – 5.66)	<0.0001	6.73 (6.22 – 7.28)

TABLE 2: Comparison of the prevalence of self-reported FA and doctor-diagnosed FA in two survey populations.

Crustacean	5.22 (4.61 – 5.92)	4.29 (3.71 – 4.94)	0.0415	4.77 (4.34 – 5.24)	4.79 (4.20 – 5.46)	2.80 (2.34 – 3.35)	<0.0001	3.83 (.344 – 4.25)
Fish	1.55 (1.23 – 1.96)	1.70 (1.35 – 2.14)	0.6097	1.62 (1.38 – 1.91)	1.37 (1.07 – 1.76)	1.10 (0.83 – 1.47)	0.2845	1.24 (1.03 – 1.50)
Mollusk	0.90 (0.66 – 1.22)	2.13 (1.73 – 2.61)	<0.0001	1.50 (1.26 – 1.78)	0.72 (0.51 – 1.01)	1.36 (1.05 – 1.76)	0.0038	1.03 (0.84 – 1.27)
Beef	0.34 (0.20 – 0.56)	2.32 (1.91 – 2.82)	<0.0001	1.30 (1.08 – 1.56)	0.27 (0.15 – 0.47)	1.46 (1.14 – 1.87)	<0.0001	0.85 (0.67 – 1.06)
Milk	0.81 (0.59 – 1.12)	0.26 (0.15 – 0.47)	0.0006	0.55 (0.41 – 0.72)	0.70 (0.49 – 0.99)	0.22 (0.11 – 0.41)	0.0012	0.46 (0.34 – 0.63)
Egg	1.15 (0.87 – 1.51)	1.10 (0.83 – 1.47)	0.9187	1.13 (0.92 – 1.37)	0.95 (0.70 – 1.28)	0.74 (0.52 – 1.05)	0.3471	0.85 (0.67 – 1.06)
Wheat	0.07 (0.02-0.20)	0.50 (0.33 – 0.77)	0.0001	0.28 (0.19 – 0.41)	0.07 (0.02 – 0.20)	0.38 (0.24 – 0.62)	0.002	0.22 (0.14 – 0.34)
Peanut	0.47 (0.31 – 0.72)	0.36 (0.22 – 0.59)	0.5046	0.42 (0.30 – 0.58)	0.27 (0.15 – 0.47)	0.31 (0.18 – 0.53)	0.7226	0.29 (0.20 – 0.43)
Soy	0.18 (0.09 – 0.35)	0.26 (0.15 – 0.47)	0.4934	0.22 (0.14 – 0.34)	0.16 (0.08 – 0.32)	0.17 (0.08 – 0.35)	>0.9999	0.16 (0.10 – 0.27)
Tree nuts	0.07 (0.02 – 0.20)	0.43 (0.27 – 0.68)	0.0006	0.24 (0.16 – 0.37)	0.02 (0.00 – 0.13)	0.31 (0.18 – 0.53)	0.0007	0.16 (0.10 – 0.27)
Other foods	0.41 (0.26 – 0.64)	0.53 (0.35 – 0.80)	0.4314	0.46 (0.34 – 0.63)	0.25 (0.14 – 0.44)	0.29 (0.16 – 0.50)	0.8354	0.27 (0.18 – 0.40)

All data were analyzed using GraphPad Prism for Windows. The Wilson/Brown method was used to calculate the 95% CIs. Fisher's exact test and Chi-square test were used (where appropriate) to compare the prevalence in two study groups. A *P* value of <0.05 was denoted as statistical significance (highlighted in bold).

TABLE 3. Multivariable logistic regression analysis of demographic factors for FA.

Risk factor	OR	P value
Gender (<i>Female/ Male</i>)	1.567	0.172
Family history of FA (<i>Yes/ No</i>)	1.018	0.961
Co-existing other allergic diseases (<i>Yes/ No</i>)	3.428	0.006
Participant location (Hue/ Tien Giang)	3.902	<0.001

Binary logistic regression was performed in SPSS Statistics for Windows to generate ORs. A *P* value of <0.05 was considered as statistical significance.

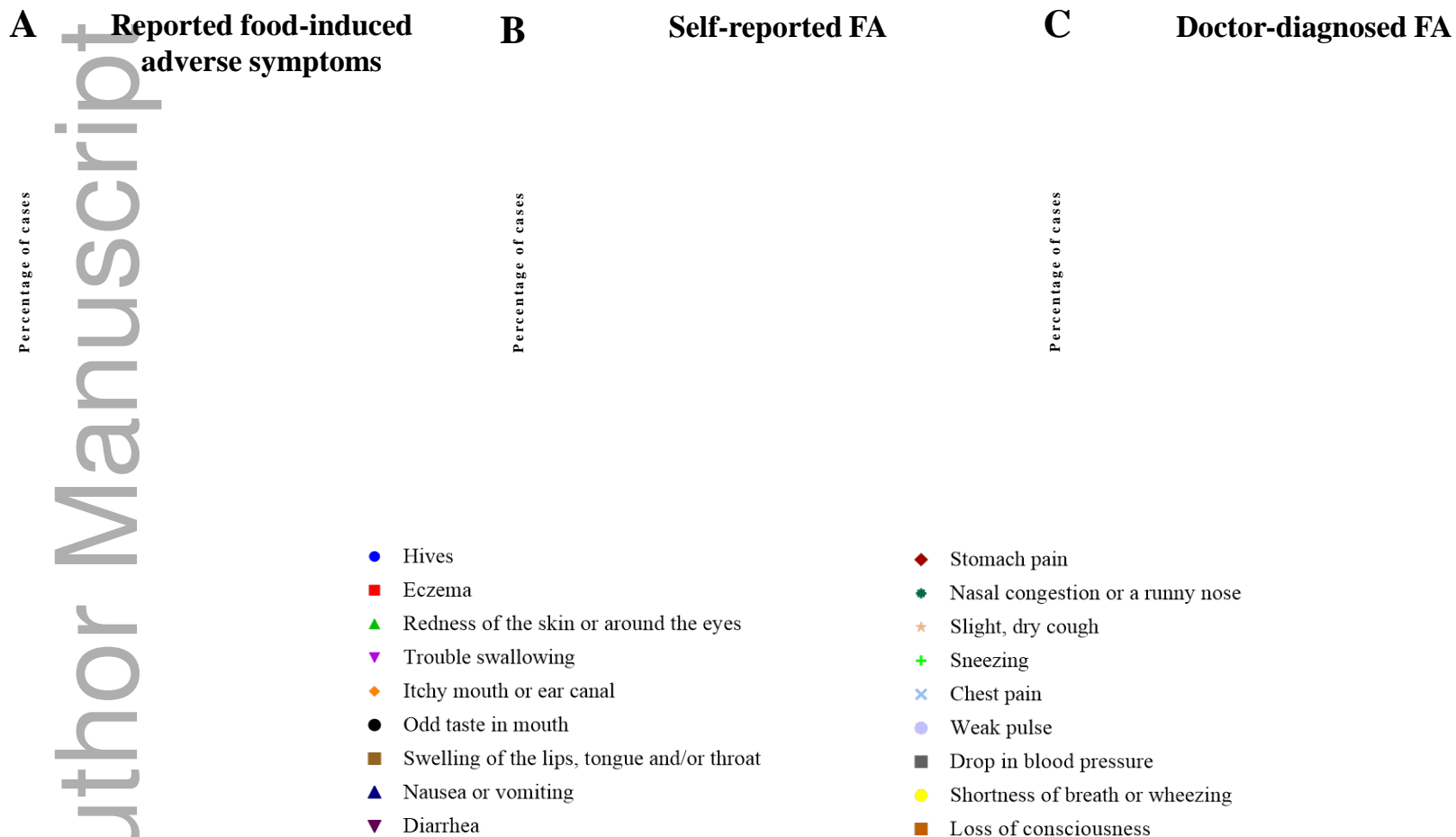


FIGURE 1. Proportion of reported clinical symptoms in participating children in Hue and Tien Giang. **A.** Reported food-induced adverse symptoms ($n = 2,905$). **B.** Reported adverse symptoms in self-reported FA participants ($n = 763$). **C.** Reported adverse symptoms in doctor-diagnosed FA participants ($n = 580$).

	Reported food-induced adverse symptoms	Self-reported FA	Doctor-diagnosed FA
Hue	A <i>n</i> = 911 Total = 1,073	B <i>n</i> = 433 Total = 496	C <i>n</i> = 373 Total = 425
Tien Giang	D <i>n</i> = 1,994 Total = 2,192	E <i>n</i> = 330 Total = 580	F <i>n</i> = 207 Total = 382

- Crustacean
- Fish
- Mollusk
- Beef
- Milk
- Egg
- Wheat
- Peanut
- Soy
- Tree nuts
- Other foods

FIGURE 2. Comparison of the distribution of reported food groups eliciting clinical reactions in participating children in Hue and Tien Giang: **A.** Reported food-induced adverse symptoms in Hue (number of participants *n* = 911); **B.** Self-reported FA in Hue; **C.** Doctor-diagnosed FA in Hue; **D.** Reported food-induced adverse symptoms in Tien Giang; **E.** Self-reported FA in Tien Giang; **F.** Doctor-diagnosed FA in Tien Giang. The total number of reported food groups is presented for each study area and symptom group.

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