The intake assessment of diverse dietary patterns on childhood hypertension: alleviating the blood pressure and lipidemic factors with low-sodium seafood rich in omega-3 fatty acids

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Abstract

Background: Childhood hypertension (CH) is related to the dietary intake and diversity of children. The study aimed to assess the critical role of dietary diversity, and seafood long-chain n-3 polyunsaturated fatty acids (LC n-3 PUFAs) in reducing CH among the Iranian community.

Methods: A cross-sectional two-phase study with 7–12-year-old Iranian students was designed. In the initial phase, the socio-demographic characteristics, and blood pressure status (normal, pre-hypertension, and hypertension) based on systolic (SBP) and diastolic (DBP) blood pressure data were assessed. The 24-h dietary recall questionnaire was used to generate the dietary diversity score (DDS, count of consumed food groups) and dietary variety score (DVS, the cumulative number of daily consumed food items). In the second phase, the association between CH reduction and changes in serum 25-hydroxyvitamin D (25OHD), total cholesterol (TC), high-density lipoprotein (HDL), low-density lipoprotein (LDL), and triglycerides (TG) levels of schoolchildren intervened by a seafood diet rich in omega-3 fatty acids were assessed using the regression analyses.

Results: The pre-hypertension and hypertension prevalence rates were 7.8 and 9.15%, respectively. CH was significantly associated with age, gender, and DDS. A significant inverse association was found between the high intake of seafood and CH ($p = 0.032$). The gas-chromatography analysis showed the high presence of α-linolenic (ALA, 6.72%), eicosapentaenoic (EPA, 7.62%), docosapentaenoic (DPA, 5.88%), and docosahexaenoic (DHA, 18.52%) acids in the seafood-based diet ($p < 0.05$). The low blood pressure levels with regular consumption of this healthy-functional diet were significantly associated with a reduction in BMI, LDL, TC, and TG, and a remarkable increase in 25OHD and HDL levels. The multiple linear regression showed that the SBP was highly associated with the TC ($p < 0.001$; $\beta = 0.464$).

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Conclusions: The age and DDS were efficient predictors for the different CH status. A regular seafood-rich dietary pattern due to the high LC n-3 PUFAs contents could significantly reduce the obesity-related cardiovascular risk factors.

Keywords: Children, Pediatric hypertension, Diet, Nutrients, Omega-3 fatty acids, Vitamin D

Introduction
Childhood hypertension (CH) is a growing health problem with a prevalence of pre-hypertension (Pre-HTN, ~3.4%) and hypertension (HTN, ~3.6%) among pediatrics aged 3 and 18 years [1]. This disorder is much higher in obese children as the Pre-HTN prevalence in obese boys and girls was reported to be 10 and 10.7%, respectively [2]. However, the CH prevalence in different countries and even in various regions of each country may be dissimilar due to the diverse genetics and environmental factors [3, 4]. The most important environmental factors affecting the CH prevalence mainly are socioeconomic status and lifestyle behaviors (e.g., dietary pattern and physical activity) [5]. Low socioeconomic status and air pollution are usually related to distress, mental health problems (such as stress, anxiety, and depression), and health-imparing behaviors [5, 6]. It was demonstrated that there was a high association between childhood hypertension and cognitive dysfunction [7]. Also, the dietary intake of foods containing high fat and sugar, as well as the low level of habitual physical activity in children can potentially increase the risk of developing hypertension during childhood [8]. The blood pressure status is determined through a balance between cardiac output (CO) and vascular resistance (VR). Therefore, an increase in each of these factors (CO or VR) could lead to an enhancement in blood pressure level as a result of changing the homeostasis of body electrolytes such as sodium, calcium, and potassium [9]. CH can occur primarily or secondary. Generally, children’s age and high blood pressure (HBP) were likely to suggest a secondary disorder. The underlying diseases should be initially controlled to treat secondary disorders [10]. Studies revealed that HBP in children could highly result in HTN disorder, cardiovascular diseases (CVDs), stroke, and even early mortality in adulthood [11]. The available evidence also proved that the structural damages to organs, such as thickening the left ventricular wall and pathological changes in vascular walls, are intensified among children with HBP levels [12]. The primarily HBP in these children was associated with other risk factors for CVDs, such as diabetes and hyperlipidemia [11, 13]. Therefore, increasing the awareness and understanding of CH risk factors should be considered as a necessity for the prevention, diagnosis, and treatment of critical underlying diseases.

Childhood and adolescence are crucial life stages to establish individual-related and health behaviors. Behavioral patterns of people in this age range were notably affected their nutritional and health status in the future [14]. Different nutritional indices have been recently introduced to assess the nutritional quality of diets and their role in the prevalence/prevention of many diseases among children and adolescents. Dietary diversity score (DDS) is a valuable indicator of the overall nutritional status. It refers to the nutritional adequacy and dietary diversity so that a higher DDS indicates a more protective effect against CVDs [15]. It has been shown that there was a strong correlation between dietary diversity and the arterial wall index, as well as the prevalence of HBP and obesity [15, 16]. The dietary diversity assessment in children to reduce HTN complications should be critically conducted. There are a few available data to support reliable conclusions related to the fruitful role of diet diversity and functional ingredients in reducing CH. Seafood is one of the most important food groups containing a high number of nutrients with remarkable benefits in human health. This food group contains n-3 polyunsaturated long chain fatty acids (e.g., eicosapentaenoic acid (EPA) and docosahexaenoic acid (DHA)), essential minerals (e.g., potassium, iodine, and selenium), as well as fat- (D, A, and E) and water-soluble vitamins (e.g., B12) [17, 18]. The weekly intake of seafood products could well reduce the postprandial triglycerides (TGs), waist circumference, and blood pressure levels, while increased the high-density lipoprotein (HDL) levels [19]. The seafood-based diet was able to maintain cognitive and antidiabetic functions by reducing inflammation [20, 21]. A reduction in the risk of ischemic stroke, congestive heart failure, coronary heart disease, and sudden cardiac death was also observed after the intake of seafoods [22–24]. Hence, the current study aimed to evaluate the relationship between dietary diversity and CH and the nutritional role of some bioactive seafood constitutes in the prevention of HTN among Iranian children aged 7–12 years.

Methods
This study was set up in two research phases (Fig. 1), including (i) the association assessment between dietary diversity and CH, and (ii) the effect
evaluation of the main bioactives of the superior nutritional diet in reducing CH.

Phase I studies

Study design and participants
In the first phase, a cross-sectional study between November 2015 and May 2016 with 514 students aged 7–12-years (274 girls and 240 boys) was conducted to investigate the effect of dietary habits on the CH prevalence. Students were chosen among Tehranian children registered in local schools of the 17th district to avoid probable sampling errors. A random number table (RNT) was used to select students after obtaining legal permissions and written informed consent. A two-stage cluster sampling in the first phase was implemented. The first and second sampling stages were the selection of primary schools and elementary students from the first to the sixth grade, respectively.

Inclusion and exclusion criteria
Inclusion criteria in the current study were schoolchildren aged 7–12 years old with Iranian nationality. Also, students with specific or chronic diseases such as diabetes, CVDs, kidney and liver diseases, cancer types, food allergies, thyroid problems, epilepsy, and seizure disorders, regular use of any medication or supplement, and unwillingness to continue cooperation with the research program were excluded. Moreover, there was no parental consanguinity in all cases.

Socio-demographic and anthropometric data
A face-to-face interview method was used to collect the data of socio-demographic characteristics of students, including age, gender, body mass index (BMI), family's economic status, and parents’ education level. The height was measured without shoes in a standing position by a stadiometer to the nearest 0.1 cm, whereas the shoulders were in a normal state. A standard, calibrated, electronic balance scale (Seca model 812, Vogel & Halke, Hamburg, Germany) with an accuracy of 0.01 g was used to assess the body weight of children. The BMI was calculated by measuring the weight (in Kg) divided by the square of assessed height (in m).

Blood pressure measurement
Systolic (SBP) and diastolic (DBP) blood pressures in the sitting position after a 5 min-rest were determined using a standard mercury pressure gauge with dimensions suitable for children and standard medical devices. In the end, the blood pressure level was divided into three groups: HTN (SBP and/or DBP ≥ 95th percentile +12 mmHg, or ≥ 140/90 mmHg), elevated or Pre-HTN (SBP and/or DBP ≥ 90th percentile but < 95th percentile, or 120/80 mmHg to <95th percentile), and normal (SBP and DBP < 90th percentile or < 120/80 mmHg) [25, 26].

Dietary intake assessment
Four trained fieldworkers during lunchtime in schools observed enrolled students and recorded the selected foods, consumed food portions, spilled or exchanged foods and leftovers on dishes in standard forms. They were entirely familiar with the type and preparation method of foods served in each school, including ten various lunch menus along with seasonal fruits and vegetables and also four items of drinks and beverages. An interviewer-administered 24-h diet recall questionnaire was also completed through face-to-face interviews in a randomized order. Two dietitians as interviewers participated in this study. The questionnaire included a list of questions related to food details and quantities (e.g., consumed portion size, ingredients of used foods, and their cooking methods) in the past 24 h [27]. An adult family member (especially the student’s mother) often helped to fill out the questionnaire for the subjects.
Determination of dietary diversity score

All the subjects completed a standard dietary diversity questionnaire according to the Food and Agriculture Organization (FAO) guidelines [28]. The diet quality was assessed using the FAO order, based on dietary variety score (DVS) to the cumulative number of various foods used in a day. The standard FAO included nine main food groups: (a) starchy materials and cereals, (b) vitamin A-rich fruits and vegetables, (c) dark green leafy vegetables, (d) other vegetables and fruits, (e) grains, nuts and legumes, dairy products, (f) organ meat (heart, liver, kidney, gizzard, and lung), (g) red meat, fish flesh, and seafoods (h) eggs, and (i) milk and dairy products. The DDS was estimated according to the minimum consumption of at least half serving of one food stated in each group. The DDS also was the total sum of the score of all food groups. Since a score of 1.0 was rated score for each group of consumed foods, the maximum score was 9.0. In this study, the 24-h recall method was used to categorize food groups using the Kant method [29]. In this methodology, the number of food groups out of five main groups with 23 subgroups as DDS was investigated as follows (Table 1):

Table 1 A summary of 24-h dietary recall questionnaire data in the studied schoolchildren in normal, Pre-HTN, and HTN groups

| Food group(s)       | Food items (preparation method)                                                                 | Normal (n = 429) | Pre-HTN (n = 47) | HTN (n = 40)  |
|---------------------|-------------------------------------------------------------------------------------------------|------------------|------------------|---------------|
|                     | Rconsuming (n)                                                                                   | MDI (g) = Energy (cal) | Rconsuming (n) | MDI (g) = Energy (cal) | Rconsuming (n) | MDI (g) = Energy (cal) |
| Bread and grains    | refined cereals (e.g., cake, white bread, pasta, biscuits, refined grain breakfast cereals, white rice, and pancakes), and whole cereal grains (e.g., wheat, corn, and rice) | 429 325 ± 28 630 ± 72 47 | 318 ± 32 603 ± 82 40 | 320 ± 41 612 ± 35 |
| Vegetables          | salad vegetables (e.g., broccoli, cabbage, cauliflower, mushrooms, bell peppers, carrots, celery, cucumbers, lettuce, peas, spinach and tomatoes), and cooked/fried potatoes | 297 214 ± 12 138 ± 27 32 | 224 ± 22 145 ± 41 28 | 215 ± 17 140 ± 29 |
| Fruits              | berries (e.g., strawberries, raspberries, blueberries, blackberries, and white and red currants), banana, citrus fruits (e.g., orange, tangerine, kiwi, and limon), apple, and fruit juices | 298 337 ± 15 196 ± 52 32 | 341 ± 21 241 ± 27 25 | 334 ± 18 232 ± 10 |
| Milk and dairy      | cow milk, cream, butter, Feta cheese, yogurt, Doogh, and Kashk                                  | 110 422 ± 75 643 ± 36 8 | 412 ± 74 654 ± 45 10 | 420 ± 45 635 ± 33 |
| products            | red meat (cooked/grilled), poultry meat (cooked/fried), and egg (boiled/pan-fried)                | 327 167 ± 11 240 ± 15 37 | 172 ± 13 254 ± 26 32 | 185 ± 9 267 ± 20 |
| Seafoods            | fried/grilled fish (e.g., salmon, trout and tuna, and carp), and fried shrimp                    | 112 376 ± 28 472 ± 51 20 | 291 ± 19 318 ± 37 8 | 252 ± 32 248 ± 28 |
| Junk foods          | salted snacks, fried fast foods, sugary carbonated beverages, gum, candy, and sweet desserts    | 164 175 ± 8 342 ± 14 18 | 180 ± 6 351 ± 14 13 | 154 ± 11 302 ± 16 |

* Rconsuming (n): Respondents with consumption (number), MDI: Mean daily intake (gram per person consuming these foods)
pattern, probably due to their high accessibility in the market. Overall, fish fillets were processed with different salt levels to increase their shelf life during the cold storage. Accordingly, the unwrapped fresh fish fillets without any added salt as an agent increasing the blood pressure were bought from the aquaculture centers in Tehran. The subjects were selected with a non-random and available sampling method from the primary population to assess the effect of the main bioactive nutrients present in this food group on the CH.

Chemicals and reagents
All the used chemicals in the analysis of fatty acids were purchased from Merck Chemical Co. (Darmstadt, Germany).

Fatty acids analysis of seafood-based diet
The method of Joseph and Ackman [30] with slight modifications was used to analyze fatty acids of recommended low-sodium seafood-based diet. In brief, the seafood oil was extracted in a Soxhlet apparatus using the extractant of petroleum ether. About 25.0 mg of oil in a 20 mL capped centrifuge tube was mixed with 1.5 mL of 0.5 M NaOH in methanol, heated in a water bath (100 °C, 5 min), and cooled at 22 ± 1 °C. Two milliliters of 12% boron trifluoride in methanol was added to the prepared mixture, re-heated (100 °C, 30 min), and then cooled before adding 1.0 mL of isoctane. In the next step, the solution was strongly stirred for 1.0 min before mixing with the saturated solution of NaCl (5.0 mL) to increase the phase separation rate. The esterified mixture was placed in the refrigerator to separate two phases, and a syringe then removed the supernatant. A mixture containing 10.0 mL of isoctane and 0.05% BHT (as an antioxidant agent) was transferred to the remaining phase into the tube, stirred to collect the formed supernatant. The sample concentrated up to 1.0 mL was then injected into the SP-2560 capillary column (100 m length × 250 μm internal diameter, 0.2 μm of film) of gas chromatography coupled with flame ionization detector (FID) (Varian 3400; Varian, Palo Alto, CA, USA). The temperature program of the column ramp was set up in the following steps: (i) the initial temperature was 70 °C, (ii) heated to 140 °C (20 °C/min) and held for 1 min; (iii) then to 180 °C (4 °C/min) and held for 1 min; and (iv) then to 225 °C (3 °C/min) and held for 30 min. The gasifying temperature was 250 °C. Nitrogen was used as the carrier gas at a flow rate of 1.0 mL/min. The injection volume was 1.0 μL in a split mode of injection at a ratio of 45:1. A comparison between the retention time of the fatty acid methyl esters (FAMEs) and their authentic standard mixtures was used to identify fatty acids. Also, the area under the peak data was used to calculate the contribution of each fatty acid as a percentage of total fatty acids using the Varian Star Chromatography Software (Varian Star WS 5.31; Varian Inc., CA, USA) [31].

Assessment of serum 25-hydroxyvitamin D
After a 12 h fasting period, antecubital venous blood samples were collected from all of the selected schoolchildren at 7:00–8:00 a.m. Serum levels of 25-hydroxyvitamin D (25OHD, ng/mL) were measured according to the procedure described by Fasihpour et al. [32] with minor modifications. In brief, the sera separation was performed by centrifugation at 3500 rpm for 25 min and then transferred into labeled polypropylene tubes to be stored at −20 °C until the analysis time of 25OHD using a commercially available enzyme-linked immunosorbent assay (ELISA) kit (DiaSource, Belgium).

Measurement of lipemic factors
The antecubital-venous blood samples of all the students were taken after an overnight fast (minimum 12 h). The total cholesterol (TC), HDL, low-density lipoprotein (LDL), and TG were measured by enzymatic methods using an Elan 2000 autoanalyzer (Eppendorf Co., Frankfurt, Germany) and expressed as mg/dL.

Statistical analysis
The obtained data were analyzed using SPSS 22.0 software (SPSS, Inc., Chicago, IL, USA). The analysis of variance (ANOVA) was employed to assess the significant quantitative differences among the studied data. The one-sample Kolmogorov-Smirnov test was used to evaluate the normality of variables’ distribution. Data were examined at a significance level of p < 0.05. Pearson’s test was also carried out to find any correlation coefficient between tested variables. Associations between SBP and factors of serum 25OHD, DBP, TC, TG, LDL, and HDL were also assessed using the multiple linear regression analysis (LRA) in multivariate adjustment with two possible confounders; i.e., age and BMI.

Results
Phase I-related findings
The prevalence of Pre-HTN and HTN in students aged 7–12 years old was determined to be 9.1 and 7.8%, respectively. The CH also was significantly correlated with the gender (p < 0.001), so that the more Pre-HTN and HTN rate in boys compared to girls was observed (Fig. 2 and b). There was an association between students’ age and high risk of HTN disorders (p < 0.001, Fig. 3). The students aged 9.5 or more showed higher rate of Pre-HTN (78.7%) and HTN (60.0%) (p = 0.001, Fig. 2c and d). Not only a considerable prevalence rate of Pre-HTN (42.6%) was detected for students aged 10 years old, but also children’s 12-year-old age group showed a high Pre-
HTN (23.3%) and HTN (25.0%) rate (Fig. 2c). The lowest and highest rates of HTN were found in the age groups of 7 and 12 years, respectively (Fig. 3). The mean SBP and DBP levels for children aged 7 years in normal, Pre-HTN, and HTN groups were 90.3 and 55.4 mmHg, 120.5 and 80.4 mmHg, and 140.5 and 90.5 mmHg, respectively. The corresponding values for children aged 12 years were 96.1 and 59.8 mmHg, 128.7 and 86.5 mmHg, and 148.8 and 94.3 mmHg, respectively. Therefore, there were significant differences in levels of SBP and DBP among schoolchildren in the different age groups (\( p < 0.05 \), Fig. 3). Moreover, the minimum (7.5%) and maximum (25.0%) number of hypertensive cases belonged to the children aged 7 and 12 years, respectively (Fig. 3).

Table 1 shows the consumption of food groups, including fruits, vegetables, red meats, dairy products, junk foods, and seafoods in normal, Pre-HTN, and HTN children groups. The amounts of mean daily intake (MDI) and received energy of each food group were also specified (Table 1). All the children in three groups of normal, Pre-HTN, and HTN consumed grains and breads. However, the consumption of other food groups in the triple groups was different. The MDI levels of breads/grains, vegetables, fruits, egg/meat products, dairy products, seafoods, and junk foods in the different groups were 318–325, 214–224, 334–341, 167–185, 412–422, 252–376, and 154–180 g, respectively (Table 1). The total absorbed calories by schoolchildren in normal, Pre-HTN, and HTN groups were 2661, 2566, and 2436 cal, respectively. Moreover, Fig. 4 depicts the association between the CH prevalence and the consumption of food groups, in normal, Pre-HTN, and HTN children groups. There was no significant relationship between the CH reduction and the consumption of red meats (\( p = 0.769 \)), dairy products (\( p = 0.430 \)), fruits and vegetables (\( p = 0.980 \)), as well as junk foods (\( p = 0.772 \)) (Fig. 4a-d). However, the consumption of seafoods among different food groups had a significant association with the prevalence reduction of CH (\( p = 0.032 \); Fig. 4e).

Figure 5 illustrates the association between the DDS and CH prevalence among the different children groups. The DDS value was significantly different in normal, Pre-HTN, and HTN children groups (\( p < 0.004 \)). There was a significant association between the DDS value (especially, \( \geq 7.0 \)) and the reduction of CH prevalence (\( p = 0.004 \)). A notable increase in CH prevalence in normal and HTN groups at a DDS of 4.0 was found. Nonetheless, the maximum CH prevalence for children in the Pre-HTN group was identified at a DDS of 6.0 (Fig. 5).

**Phase II related-findings**

The lipid content of the seafood diet was 13.72%. The GC analysis revealed that the main fatty acids present in
seafood diet were linoleic acid (18:2; 3.65%), α-linolenic acid (ALA, 18:3; 6.72%), EPA (20:5; 7.62%), docosapentaenoic acid (DPA, 22:5; 5.88%), and DHA (22:6; 18.52%). There was a significant difference in the content of fatty acids present in the seafood diet (p < 0.05). Table 2 showed that the low-sodium seafood-based dietary treatment could significantly reduce BMI, SBP, DBP, LDL, TG, and TC levels in boys, girls, and the total population (p < 0.05). Also, this nutritional intervention increased the HDL and 25OHD levels in all the populations significantly (p < 0.05). There was no significant difference in BMI, 25OHD, blood pressure, and lipidemic factors after the intervention among boys, girls, and total populations (Table 2). However, an insignificantly higher 25OHD, SBP, LDL, TG, and TC values in boys compared to girls were observed (Table 2).

Table 3 reveals the results of the Pearson’s correlation test between age, BMI, blood pressure (i.e., SBP and DBP), lipid profile (i.e., LDL, DHL, TG, and TC), and vitamin D amounts in the study population. A highly significant correlation was explored among all the investigated parameters. Nonetheless, there was no significant association between serum 25OHD and age. The maximum correlation was detected between LDL and HDL (p < 0.001; r = −0.936). Conversely, the lowest significant correlation rate was identified between LDL/HDL and

![Fig. 3](image-url)
age, as well as 25OHD and SBP. In general, a negative, significant correlation was found between 25OHD/HDL levels and other parameters (i.e., age, BMI, DBP, SBP, LDL, TG, and TC). Table 4 also exhibits the multiple linear regression results of all the variables as a function of SBP. According to the fitted model ($R^2 = 0.825$; adjusted $R^2 = 0.794$), the effect of gender, DBP, HDL, and TC on the SBP changes was highly significant. Among these variables, TC had the highest role in SBP changes. For every 1 mg/dL increase in TC, the SBP was expected to rise by 0.464 mmHg. However, the LDL, BMI, TG, age, and 25OHD were not significantly associated with the SBP.

**Discussion**

The prevalence of Pre-HTN and HTN among 12–7-year aged students was reported to be 9.1 and 7.8%, respectively. The similar prevalence rate for CH in other cities...
Table 2 A comparative study on levels of BMI, blood pressure, serum vitamin D, and lipidemic factors among girls (n = 29), boys (n = 31), and total population (n = 60) before and after the low-sodium seafood-based dietary intervention

| Parameter | Boys Before | After | Girls Before | After | Total Before | After |
|-----------|-------------|-------|--------------|-------|--------------|-------|
| BMI (kg/m²) | 17.53 ± 0.32<sup>a</sup> | 17.07 ± 0.21<sup>a</sup> | 17.25 ± 0.18<sup>a</sup> | 17.00 ± 0.22<sup>a</sup> | 17.41 ± 0.19<sup>a</sup> | 17.04 ± 0.21<sup>a</sup> |
| 25OHD (ng/mL) | 17.89 ± 3.54<sup>b</sup> | 20.49 ± 5.34<sup>a</sup> | 16.77 ± 3.49<sup>b</sup> | 18.37 ± 3.84<sup>a</sup> | 17.74 ± 3.95<sup>b</sup> | 19.46 ± 4.76<sup>a</sup> |
| SBP (mmHg) | 106.71 ± 0.26<sup>a</sup> | 102.64 ± 0.68<sup>b</sup> | 105.41 ± 0.22<sup>a</sup> | 102.02 ± 1.03<sup>b</sup> | 106.02 ± 0.31<sup>a</sup> | 102.34 ± 0.91<sup>b</sup> |
| DBP (mmHg) | 62.37 ± 0.08<sup>a</sup> | 60.92 ± 0.52<sup>b</sup> | 62.97 ± 0.34<sup>a</sup> | 60.93 ± 0.57<sup>b</sup> | 62.54 ± 0.27<sup>a</sup> | 60.92 ± 0.54<sup>b</sup> |
| LDL (mg/dL) | 109.54 ± 0.85<sup>a</sup> | 99.92 ± 4.71<sup>b</sup> | 107.42 ± 0.61<sup>a</sup> | 98.70 ± 5.72<sup>b</sup> | 108.05 ± 0.74<sup>a</sup> | 99.33 ± 5.21<sup>b</sup> |
| HDL (mg/dL) | 32.26 ± 1.75<sup>b</sup> | 39.83 ± 5.26<sup>a</sup> | 31.75 ± 1.57<sup>b</sup> | 40.20 ± 6.11<sup>a</sup> | 32.09 ± 1.28<sup>b</sup> | 40.01 ± 5.64<sup>a</sup> |
| TG (mg/dL) | 88.25 ± 4.04<sup>a</sup> | 76.09 ± 7.43<sup>b</sup> | 83.32 ± 3.77<sup>a</sup> | 75.80 ± 9.30<sup>b</sup> | 84.65 ± 2.98<sup>a</sup> | 75.95 ± 8.31<sup>b</sup> |
| TC (mg/dL) | 182.32 ± 2.65<sup>a</sup> | 169.83 ± 7.73<sup>b</sup> | 184.23 ± 2.24<sup>a</sup> | 169.17 ± 9.76<sup>b</sup> | 181.29 ± 1.89<sup>a</sup> | 169.51 ± 8.70<sup>b</sup> |

<sup>a</sup> The mean age of boys, girls, and total population was 9.02 ± 1.60, 9.31 ± 1.47, and 9.16 ± 1.53 years, respectively

<sup>b</sup> The means (in each row) for each children's group with dissimilar letters (a,b) are significantly different (p < 0.05)

of Iran for the same age groups was reported. Mehr-Alizadeh et al. [33] reported an HTN prevalence rate of 3–9.6% for 7–17-year aged students in Semnan city. Also, Zardast et al. [34] estimated a prevalence rate of 11.6 and 7.4% for Pre-HTN and HTN in 6–11 year aged children in Birjand city, respectively. It seems that the economic situation was critical in CH development. Since the 17th district located in the southern part of Tehran had a similar economic situation to most cities of Iran, the results obtained in Semnan and Birjand cities were in line with our findings. The age elevation was associated with an increased HTN risk. Similarly, some researchers found that higher age was related to a rise in the CH prevalence [35, 36]. Patil and Garg [37] and Hakim et al. [38] also mentioned that the children's age was associated with their average SBP and DBP. An integration between nutritional diet and physical activity pattern in students 7 years old might prevent the prevalence of Pre-HTN. It seems that Iranian parents in the first year of school had serious attention to their children's diet and exercise. However, reasons for higher pre-HTN prevalence at ages more than 7 years old might be related to lower attention of parents to the accurate lifestyle of students by increasing their age in terms of

Table 3 Correlation study between age, BMI, blood pressure, lipid profile and vitamin D levels in the study population

| Pearson' test | Age (yrs. old) | BMI (kg/m²) | SBP (mmHg) | DBP (mmHg) | LDL (mg/dL) | HDL (mg/dL) | TG (mg/dL) | TC (mg/dL) | 25OHD (ng/mL) |
|---------------|----------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|---------------|
| Age (yrs. old) | 1              | r = 0.53     | r = 0.45    | r = 0.53    | r = 0.30    | r = −0.28   | r = 0.719   | r = 0.406   | r = −0.252    |
| BMI (kg/m²)     | r = 0.553      | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.019   | p = 0.028   | p < 0.001   | p = 0.001   | p = 0.052<sup>ns</sup> |
| SBP (mmHg)      | r = 0.452      | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p = 0.008     |
| DBP (mmHg)      | r = 0.528      | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p = 0.004     |
| LDL (mg/dL)     | r = 0.303      | p = 0.019   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p = 0.004     |
| HDL (mg/dL)     | r = −0.283     | p = 0.028   | p < 0.001   | p < 0.001   | p < 0.001   | p = 0.001   | p < 0.001   | p < 0.001   | p = 0.001     |
| TG (mg/dL)      | r = 0.719      | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001     |
| TC (mg/dL)      | r = 0.406      | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001     |
| 25OHD (ng/mL)   | r = −0.252     | p = 0.008   | p = 0.022   | p = 0.004   | p < 0.001   | p < 0.001   | p < 0.001   | p < 0.001   | p = 0.010     |

<sup>ns</sup> non-significant
high-fat diets and lower physical activities. As a result, this increase may be owing to the increase in body mass, which occurs as the child grows up. However, this fact in adult populations was attributed to the function reduction of the body’s antioxidant enzymes (e.g., superoxide dismutase) by aging. The HTN was also increased with a decrease in the synthesis of nitric oxide as the body’s vasodilator [39]. The significantly higher HTN prevalence in boys than girls was found. This was consistent with the study of Brady et al. [8] in the US, who studied the HTN prevalence among 184 children aged 3–20 years. However, the significantly higher prevalence of HTN in girls compared to boys in Shiraz, Iran, was earlier reported by Ayatollahi and Zare [40]. The difference in lifestyle, diet patterns, and physical activities might be involved in the different prevalence of HTN in girls and boys before puberty.

According to the received energy data, the calorie intake levels in the 24-h recall method was more than the recommended calorie levels of the DASH diet. This result may be due to the higher consumption of junk foods and the lower number of seafood-related servings. Based on the dietary recommendations of therapeutic lifestyle changes (TLC) and dietary approaches to stop hypertension (DASH), receiving the food groups of fruits and vegetables (high content of potassium, magnesium, stanol, sterol, and fibers), as well as dairy products (remarkable amounts of calcium and vitamin D) had an important role in reducing CH. In contrast, the meat group had a direct relationship with the CH prevalence due to the high content of sodium and saturated fatty acids (SFAs). However, we found that the consumption of diets based on fruits, vegetables, and dairy products, along with meats, had not any significant impact on the prevalence of HTN and Pre-HTN in children. It was indicated that the imbalance of mineral metabolism (i.e., calcium, potassium, and sodium) in the daily intake of food groups is a more significant factor in the HTN prevalence [41]. This result was probably owing to the low efficiency of the methodology (Kant method) used in the current study to determine the dietary groups. It was recommended to daily receive more than five servings of fruits and vegetables for reducing or controlling the HTN risk. However, only half servings per day from each child’s diet group in the current research was assigned to this food group. Besides, the increased SFAs, cholesterol, and hidden salt content in high-fat dairy products might notably eliminate their beneficial effects on the status of blood pressure. Similarly, the group of red meats might have destructive effects on the blood pressure because of the high content of sodium and SFAs. It was hypothesized that the CH could be directly escalated by consuming junk foods (e.g., chips, puffs, carbonated beverages, and sodas) as a result of high amounts of calories and sodium. Stanley et al. [42] considered the high content of sugar, fat, and sodium in junk foods as possible risk factors for HBP. Nonetheless, this theory was not demonstrated because we found that the consumption of junk foods has not any significant relationship with the occurrence of HTN and Pre-HTN in children. On the other hand, the consumption effects of seafood on CH reduction were because of the beneficial role of omega-3 fatty acids such as the vascular response to angiotensin, the improvement of vascular function, and the inflammation moderation or the attenuation of proinflammatory reactions involved in hypertension [43–45]. Seafood is also considered as a good protein source with high digestibility, bioavailability, and efficiency ratio. The presence of essential amino acids (e.g., cystine, lysine, methionine, threonine, and

### Table 4

| Model | Unstandardized coefficients | Standardized coefficients | p-value |
|-------|-----------------------------|---------------------------|---------|
|       | β                           | SE                        | t       | p-value     |
| Constant | 67.363                      | 8.434                     | 7.987   | < 0.001     |
| Age (year) | 0.074                      | 0.042                     | 0.123   | 1.778       | 0.081<sup>bc</sup> |
| BMI (kg/m<sup>2</sup>) | 0.446                      | 0.438                     | 0.106   | 1.019       | 0.313<sup>bc</sup> |
| Gender | −0.575                      | 0.108                     | −0.316  | −5.322      | < 0.001     |
| DBP (mmHg) | 0.474                      | 0.148                     | 0.280   | 3.19        | 0.002       |
| HDL (mg/dL) | −0.333                     | 0.012                     | −0.203  | −2.677      | 0.010       |
| TG (mg/dL) | 0.004                      | 0.015                     | 0.039   | 0.288       | 0.775<sup>bc</sup> |
| TC (mg/dL) | 0.049                      | 0.009                     | 0.464   | 5.248       | < 0.001     |
| 25OHD (ng/mL) | 0.015                     | 0.016                     | 0.077   | 0.914       | 0.365<sup>bc</sup> |
| Excluded variable | β                        | t                        | p-value | Partial correction | Tolerance |
| LDL (mg/dL) | −0.059                     | −0.257                    | 0.775<sup>bc</sup> | −0.041      | 0.064       |

<sup>a</sup> Multiple linear regression model: R<sup>2</sup> = 82.5%, adjusted R<sup>2</sup> = 79.4%

<sup>b</sup> p-value < 0.05 is significant, ns: non-significant
tryptophan) in the structure of angiotensin I–converting enzyme inhibitory peptides may reduce the CH [46]. The consumption of seafood products through high daily intake of dietary potassium is associated with the reduced ratio of sodium (Na) to potassium (K) and blood pressure levels in children [47]. However, the synergy effect of different nutrients of seafood sources on the CH reduction has been less investigated.

The higher dietary diversity can cause a lower calorie density of consumed foods. Even though the DDS was significantly different in normal, Pre-HTN, and HTN children, a direct or reverse relationship for this difference was not generalizable. However, most children in both groups of normal and Pre-HTN received a DDS score of 4.0. Results of this study were in line with findings reported by Lazarou et al. [48], who explored the relationship between dietary patterns and blood pressure levels using the E-KINDEX scoring system. In this methodology, the consumed foods were divided into 13 groups and individual points from their receipt. Results indicated that the E-KINDEX score of foods had been independently linked to lower blood pressure among healthy children. Also, our findings were consistent with the results of Miller et al. [49] as there was not a link between dietary diversity and the prevalence of hypertension. They explained that the imbalance exists in the dietary intake which could significantly lead to the prevalence of hypertension [49]. Obesity is one of the most critical risk factors for hypertension. It is strongly advised to reduce the weight and to receive less high-quality foods in the nutritional medicine for preventing and treating hypertension in children. The higher diet diversity could cause a lower calorie density of consumed food [50]. However, despite the significant difference in DDS value on the status of blood pressure in children, there was no linear relationship among normal, pre-hypertension and hypertension children. Thus, higher or lower DDS did not necessarily mean an association with the hypertension status of children. As a result, it was suggested that not only the scoring method used by the FAO should be implemented, but also the dietary quality score (DQS) should be calculated.

The intake of a low-sodium seafood-based diet could significantly reduce the levels of BMI, blood pressure, and adverse lipidemic factors (such as LDL, TG, and TC) in the study population. In contrast, the serum vitamin D and HDL levels were remarkably increased after implementing the seafood-based dietary intervention. Tu et al. [51] mentioned that obesity and body fat in children were directly associated with their HBP. Therefore, fish oil and omega-3 supplementation can be used as complementary therapies for BMI reduction in obese and overweight patients with HTN and CVDs. Earlier, Kodas et al. [52] in animal studies pointed out that the presence of long-chain n-3 polyunsaturated fatty acids (LC n-3 PUFAs) could modulate the function of the major brain neurotransmitters such as noradrenaline secretion, as well as serotonin and dopamine systems, which are usually involved in the adjustment of blood pressure and heart rhythm. Baltzell et al. [53] explained that the enzyme of lipoprotein lipase (LPL) located in the endothelial layer of capillaries in the muscle and adipose tissues had a high potential to hydrolyze chylomicron- and very-LDL (VLDL)-triacylglycerol to their structural fatty acids. Although the intake of LC n-3 PUFAs with the fish consumption elevated the activity of muscle LPL, the adipocyte LPL activity was decreased. Thus, the changed LPL activities were along with a reduction in body fat and plasma TG levels, showing the TG utilization was altered from storage form in adipocytes to oxidized form in muscles after the consumption of LC n-3 PUFAs [54]. In addition, the hypocholesterolemic effects of fish proteins accompanied by lower TG levels were previously reported [55, 56].

Limitations
There were some limitations to the present study. It seems that Kant’s method was inadequate to assess the influence of food groups on the levels of blood pressure in children. For example, it is suggested that the type of consumed meat will be considered in two marine and non-marine groups in future studies. The use of weekly intake as a criterion for the intake level assessment of junk foods likely was not sufficiently useful. Besides, the consumption recording of such foods in the interview possibly resulted in an increase in the response error due to the difficulty of remembering the time of day of food intake. It is recommended to use a feed registration method or a 24-h recall questionnaire in 3 days (two days a week and one weekend) to evaluate the weekly intake for junk foods. Also, there was a strict requirement for further high-quality population-based investigations and randomized controlled trials of long duration to comprehensively assess the effects of LC n-3 PUFAs and fish on renal function and its role in reducing the HTN rate.

Conclusion
This study showed that high dietary diversity could be a powerful tool in reducing the HTN prevalence rate among Iranian schoolchildren. The intake of junk foods, fruits and vegetables, red meats, and dairy products were not significantly correlated with CH. However, a significant association between seafood intake and reduced blood pressure demonstrated the nutritional efficiency of LC n-3 PUFAs in reducing the children’s blood pressure levels. The regular consumption of a healthy seafood-based diet could significantly reduce HTN and BMI due
to the high presence of ALA, EPA, DPA, and DHA, the increased 25OHD and HDL levels, and the reduced amounts of LDL, TC, and TG. The future practical nutritional programs should be settled on the effectiveness of different seafood-based dietary patterns in the prevention of CH, overweight, and obesity as risk factors for CVDs. The choice of innovative non-thermal cooking ways for the preservation of nutritional constitutes present in seafood products is necessary as it has an influential role in childhood nutrition and wellbeing. Severe attention should be paid towards the implementation of case-control studies with an extensive follow up period to evaluate long-term effects of different seafood products and their nutritional benefits. Finally, more attention in the public health framework should be directed to improve dietary diversity and preparation methods of particular food items.

Abbreviations

CH: Childhood hypertension; LC n-3 PUFAs: Long-chain n-3 polyunsaturated fatty acids; DDS: Dietary diversity score; DVS: Dietary variety score; DQS: Dietary quality score; MDI: Mean daily intake; 25OHD: 25-hydroxyvitamin D; TC: Total cholesterol; LDL: Low-density lipoprotein; HDL: High-density lipoprotein; DASH: Dietary approaches to stop hypertension; CO: Cardiac output; VR: Vascular resistance; HTN: Hypertension; CO: Cardiac output; VR: Vascular resistance; LDL: Low-density lipoprotein; HDL: High-density lipoprotein; FAMEs: Fatty acid methyl esters; EPA: Eicosapentaenoic acid; DHA: Docosahexaenoic acid; DDA: Docosapentaenoic acid; Pre-HTN: Pre-hypertension; HTN: Hypertension; CO: Cardiac output; VR: Vascular resistance; HD: High-density lipoprotein; LDL: Low-density lipoprotein; TG: Triglyceride; BMI: Body mass index; ALA: α-linolenic acid; PUFAs: Polyunsaturated fatty acids; FAMES: Fatty acid methyl esters; SYM: Symptom; 25OHD: 25-hydroxyvitamin D; TC: Total cholesterol; LDL: Low-density lipoprotein; HDL: High-density lipoprotein; VLDL: Very-low-density lipoprotein; DASH: Dietary approaches to stop hypertension; SFA: Saturated fatty acids.

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Authors’ contributions

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Availability of data and materials

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

Ethics approval and consent to participate

Informed written consent was obtained from all participants and their parents, whereas the participation was entirely voluntary and confidentiality assured. The study was also performed following the Declaration of Helsinki and approved by the Human Ethics Committee of X University of Medical Sciences.

Consent for publication

Not applicable.

Competing interests

The authors declare that they have no competing interests.

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References

1. Flynn JT, Falkner BE. New clinical practice guideline for the management of high blood pressure in children and adolescents. Hypertension. 2017;70(4):683–6.
2. Yilmaz M, Aktaç B, Dereli F, San HY, Ağartıklı G, Gündoğdu ZT. Assessment of the prevalence of obesity, stunting, and hypertension among primary school children. Eur J Ther. 2017;23(3):99–105.
3. Hu J, Shen H, Wu JZ, Xiao Q, Chu GP, Teng CG, et al. Prevalence of high blood pressure and high normal blood pressure among 7- to 17-year-old children and adolescents in developed regions, China from 2014 to 2017: using new national blood pressure reference for Chinese children and adolescents. J Hum Hypertens. 2019;33(5):400–10.
4. Ebrahimzadeh M, Emarani M, Hashemini H, Fotouhi A. Prevalence of prehypertension and hypertension and its risk factors in Iranian school children: a population-based study. J Hypertens. 2018;36(9):1816–24.
5. Feber J, Litwin M. Primary hypertension. In: Hypertension in children and adolescents. Cham: Springer; 2019. p. 95–110.
6. Tyner E, Oropeza M, Figueroa J. Childhood hypertension and effects on cognitive functions: mechanisms and future perspectives. CNS Neurol Disord-DR. 2019;18(9):677–86.
7. Katamba G, Agaba DC, Migisha R, Namaganda A, Namayanja R, Turyakira E. Using blood pressure height index to define hypertension among secondary school adolescents in southwestern Uganda. J Hum Hypertens. 2020;34(1):76–81.
8. Brady TM, Fivush B, Flynn JT, Parekh R. Ability of blood pressure to predict left ventricular hypertrophy in children with primary hypertension. J Pediatr. 2008;152(1):73–8.
9. Cheang MH, Kowalk GT, Quail MA, Steeden JA, Hothi D, Tullus K, et al. The cardiovascular phenotype of childhood hypertension: a cardiac magnetic resonance study. Pediatr Radiol. 2019;49(6):727–36.
10. Dhull RS, Baracco R, Jain A, Matteo TK. Pharmacologic treatment of pediatric hypertension. Curr Hypertens Rep. 2016;18(4):32.
11. Amiri P, Vahedi-Notash G, Naseri P, Khalili D, Nazari SS, Mehvari Y, et al. National trends of pre-hypertension and hypertension among Iranian adolescents across urban and rural areas (2007–2011). Biol Sex Differ. 2019;10(1):15.
12. Meng L, Hou D, Zhao X, Hu Y, Liang Y, Liu J, et al. Cardiovascular target organ damage could have been detected in sustained pediatric hypertension. Blood Press. 2015;24(5):284–92.
13. Vijayalakshimi I, Narasimhan C. Hypertension in children and adolescents. Hypertension. 2015;1(2):88–93.
14. Lande MB, Kupferman JC, Adams HR. Cognitive and behavioral aspects of childhood hypertension. In: Flynn JT, Ingelfinger JR, Redwine KM, editors. Pediatric hypertension. Totowa, NJ: Humana Press; 2013. pp. 465–78
15. Salehi-Abargouei A, Akbari F, Bellissimo N, Azadbalikh L. Dietary diversity score and obesity: a systematic review and meta-analysis of observational studies. Eur J Clin Nutr. 2016;70(1):1–9.
16. Farhangi MA, Jahangiry L. Dietary diversity score is associated with cardiovascular risk factors and serum adiponectin concentrations in patients with metabolic syndrome. BMC Cardiovasc Disord. 2018;18(1):68.
17. Innes JK, Calder PC. Marine omega-3 (n-3) fatty acids for cardiovascular health: an update for 2020. Int J Mol Sci. 2020;21(4):1362.
18. Hosomi R, Yoshida M, Fukunaga K. Seafood consumption and components for health. Glob J Health Sci. 2012;4(3):72–86.
19. Schmedes M, Bald exact EK, Jacques H, Lavigne C, Graff IE, et al. The effect of lean-seafood and non-seafood diets on fasting and postprandial serum metabolites and lipid species: results from a randomized crossover intervention study in healthy adults. Nutrients. 2018;10(5):E598.
40. Ayatollahi SM-T, Zare M. Blood pressure standards for Shiraz (southern Iran) Izadi
42. Stanley WC, Shah KB, Essop MF. Does junk food lead to heart failure?
41. Lau WL, Pai A, Moe SM, Giachelli CM. Direct effects of phosphate on vascular cell function. Adv Chronic Kidney Dis. 2011;18(2):105–12.
44. Dangardt F, Osika W, Chen Y, Nilsson U, Gan LM, Gronowicz E, et al. Omega-3 fatty acid supplementation improves vascular function and reduces inflammation in obese adolescents. Atherosclerosis. 2010;212(2):2580–5.
45. Yang B, Ren XL, Li ZH, Shi MQ, Ding F, Su KP, et al. Lowering effects of fish oil supplementation on proinflammatory markers in hypertension: results from a randomized controlled trial. Food Funct. 2020;11:1779–89.
46. Ghassemi M, Babji AS, Said M, Mahmodani F, Aihara K. Angiotensin I–converting enzyme inhibitory peptides from snakehead fish sarcoplasmic protein hydrolysate. J Food Biochem. 2014;38(2):140–9.
47. O’Halloran SA, Grimes CA, Lacy KE, Campbell KJ, Nowson CA. Dietary intake and sources of potassium and the relationship to dietary sodium in a sample of Australian pre-school children. Nutrients. 2016;8(8):496.
48. Lazaro C, Paragotobos DB, Matalas A-L. Foods E-KINDEX: a dietary index associated with reduced blood pressure levels among young children: the CYKIDS study. J Am Diet Assoc. 2009;109(6):1070–5.
49. Miller WL, Crabtree BF, Evans DK. Exploratory study of the relationship between hypertension and diet diversity among Saba islanders. Public Health Rep. 1992;107(4):426–52.
50. Azadbakht L, Esmailzadeh A. Dietary energy density is favorably associated with dietary diversity score among female university students in Isfahan. Nutrition. 2012;28(10):991–5.
51. Tu W, Eckert GJ, DiMeglio LA, Yu Z, Jung J, Pratt JH. Intensified effect of adiposity on blood pressure in overweight and obese children. Hypertension. 2011;58(5):818–24.
52. Das A, Calineau L, Bodard S, Vancasselt S, Guilleudeau D, Besnard JC, et al. Serotoninergic neurotransmission is affected by n-3 polyunsaturated fatty acids in the rat. J Neurochem. 2004;89(3):695–702.
53. Baltzell JK, Wooten JT, Otto DA. Lipoprotein lipase in rats fed fish oil: apparent relationship to plasma insulin levels. Lipids. 1991;26(4):289–94.
54. Bonafini S, Antoniazzi F, Maffei C, Minuz P, Fava C. Beneficial effects of ω-3 PUFAs in children on cardiovascular risk factors during childhood and adolescence. Prostaglandins Other Lipid Mediat. 2015;120:72–9.
55. Demonty I, Deshaies Y, Lamarche B, Jacques H. Cod protein lowers the hepatic triglyceride secretion rate in rats. J Nutr. 2003;133(5):1308–402.
56. Saravanan P, Davidson NC, Schmidt EB, Calder PC. Cardiovascular effects of marine omega-3 fatty acids. Lancet. 2010;376(9740):540–50.