Higher adherence to the Mediterranean Diet is associated with lower micronutrient inadequacy in children: the SENDO project

Asier Oliver Olid¹, Elise Fabios², Lorena García-Blanco³, Jose Manuel Moreno-Villares⁴, Miguel Ángel Martínez-González^{5,6,7,8} and Nerea Martín-Calvo^{5,6,7,*}

¹Department of Pediatrics, Hospital Universitario de Navarra, Pamplona, Spain: ²University of Navarra, School of Medicine, Department of Preventive Medicine and Public Health, Pamplona, Spain: ³Olite Primary Care Health Center, Servicio Navarro de Salud-Osasunbidea, Pamplona, Spain: ⁴Department of Pediatrics, Clinic Universidad de Navarra, Madrid, Spain: ⁵University of Navarra, School of Medicine, Department of Preventive Medicine and Public Health, Pamplona, Spain: ⁶IdiSNA, Instituto de Investigación Sanitaria de Navarra, Pamplona, Spain: ⁷CIBER Pathophysiology of Obesity and Nutrition, Carlos III Health Institute, Madrid, Spain: ⁸Department of Nutrition, Harvard TH Chan School of Public Health, Boston, MA, USA

Submitted 5 May 2023: Final revision received 19 October 2023: Accepted 27 November 2023

Abstract

Objective: To assess whether the Mediterranean Diet (MedDiet) is associated with lower micronutrients inadequacy in a sample of Spanish preschoolers.

Design: We conducted a cross-sectional study with 4–5-year-old children participating in the SENDO project. Information was gathered through an online questionnaire completed by parents. Dietary information was collected with a previously validated semi-quantitative FFQ. The estimated average requirements or adequate intake levels as proposed by the Institute of Medicine were used as cut-off point to define inadequate intake.

Statistical analyses: Crude and multivariable adjusted estimates were calculated with generalised estimated equations to account for intra-cluster correlation between siblings.

Participants: We used baseline information of 1153 participants enrolled in the SENDO project between January 2015 and June 2022.

Main outcomes measures: OR and 95% CI of presenting an inadequate intake of ≥ 3 micronutrients associated with the MedDiet.

Results: The adjusted proportion of children with inadequate intake of ≥ 3 micronutrients was 27·2 %, 13·5 % and 8·1 % in the categories of low, medium and high adherence to the MedDiet, respectively. After adjusting for all potential confounders, children who had a low adherence to the MedDiet showed a significant lower odds of inadequate intake of ≥ 3 micronutrients compared to those with a high adherence (OR 9·85; 95 % CI 3·33, 29·09).

Conclusion: Lower adherence to the MedDiet is associated with higher odds of nutritional inadequacy.

Keywords Breast-feeding Micronutrients Diet quality Children

In recent years, nutritional science has shifted from a reductionist paradigm, mainly focused on single nutrients, to a more holistic approach to diet that considers global dietary quality⁽¹⁾. Most international dietary guidelines now endorse a dietary pattern approach, moving away from quantitative nutrient advice⁽²⁾.

The 2020–2025 Dietary Guidelines for Americans define a dietary pattern as 'the quantities, proportions, variety, or combination of different foods, drinks, and nutrients in diets, and the frequency with which they are habitually consumed'⁽³⁾. This better reflects the relation between food

and health, including in the equation the complex interactions of nutrients and non-nutrients within our organism. Moreover, it presents clear advantages in clinical practice and in terms of public health messages, as it is more easily translatable to the general population than the traditional quantitative measures, that made guidelines hard to grasp and adherence low^(2,4).

Among the dietary patterns with the greatest scientific consensus, the Mediterranean Diet (MedDiet) has been recognised as one of the healthiest diets worldwide, as evidence has shown to protect against multiple chronic

*Corresponding author: Email nmartincalvo@unav.es

© The Author(s), 2023. Published by Cambridge University Press on behalf of The Nutrition Society. This is an Open Access article, distributed under the terms of the Creative Commons Attribution-NonCommercial-NoDerivatives licence (http://creativecommons.org/licenses/by-nc-nd/4.0/), which permits non-commercial re-use, distribution, and reproduction in any medium, provided that no alterations are made and the original article is properly cited. The written permission of Cambridge University Press must be obtained prior to any commercial use and/or adaptation of the article.





diseases and increase life expectancy^(5–8). The MedDiet is characterised by a high consumption of plant-based foods, moderate-to-high consumption of fish and low consumption of meat and dairy products (with the exception of vogurt and long preservable cheeses). Nevertheless, the hallmark of the MedDiet is the liberally consumed olive oil (or extra virgin olive oil), which represents its main culinary fat⁽⁹⁾. These foods make the MedDiet be rich in micronutrients, fibre, antioxidants and healthy fats and therefore a stellar candidate to achieve nutritional adequacy.

Micronutrients are vitamins and minerals involved in multiple functions such as the production of enzymes and hormones⁽¹⁰⁾. An insufficient dietary intake of micronutrients leads to a depletion of body reserves which, if maintained, is followed by a decrease in serum levels and then by clinical impairment. Although severe deficits are more frequent in low-income countries, inadequate micronutrient intake is frequent in middle- and high-income countries(11,12).

Suboptimal nutrition is a key concern in children around the world^(13,14), including Spain⁽¹⁵⁾. Evidence has shown that the MedDiet enhanced nutritional adequacy in adult populations⁽¹⁶⁾, but, to our knowledge, few studies have studied this association in children, and none in preschoolers particularly. For this reason, we investigated the association between adherence to the MedDiet and nutrient adequacy in a population of Spanish preschoolers.

Materials and methods

Study population

The Seguimiento del Niño para un Desarrollo Óptimo (SENDO) project is an ongoing Spanish prospective cohort focused on the study of the effect of diet and lifestyle on the health of children and adolescents. The recruitment is permanently open. Participants are invited to enter the cohort by their paediatrician at their primary care health centre or by the research team at school. The cohort has the following inclusion criteria: (1) children of 4 or 5 years of age and (2) residing in Spain. The sole exclusion criterion is the lack of access to an internet-connected device to complete the questionnaires. Information is collected at baseline and updated every year through self-administered online questionnaires, which are completed by parents. For this study, we used baseline information of participants recruited between January 2015 and June 2022. Of the 1153 participants in the SENDO project recruited up to June 2022, 138 were excluded for presenting energy values > p99 or < p1, 111 for presenting implausible micronutrient intake values (mean ± 3 sD) and 81 for not having completed the baseline questionnaire. A total of 819 preschoolers were finally included (Fig. 1).

The SENDO project follows the rules of the Helsinki Declaration on Ethical Principles for Human Research, and its protocol was approved by the Ethical Committee for

Clinical Research of Navarra (Pyto 2016/122). Participants' parents or legal guardians signed an informed consent before entering the study.

Assessment of the exposure

Dietary information was collected at baseline with a previously validated 147-item semi-quantitative FFQ⁽¹⁷⁾. A portion size was specified for each food item. Parents reported how often their child had consumed each of the food items over the previous year by choosing one out of the nine options of response ranging between 'never/ almost never' and '≥ 6 times/day'. The nutrient content of each food item was calculated by trained dietitians, by multiplying the frequency of consumption by the edible portion and the nutrient composition of the specified portion size. Updated Spanish food composition tables⁽¹⁸⁾ and online information⁽¹⁹⁾ were used for this purpose. Total energy intake was obtained by adding the calorie contribution of each item.

Diet quality was assessed with the KIDMED index, an a priori-defined dietary index to evaluate the adherence to the MedDiet pattern in children and adolescents⁽²⁰⁾. The KIDMED index consists of sixteen items, of which twelve items score 0 or +1 and four items score -1 or 0. Thus, the score in the KIDMED index may range from -4 to 12 points. Participants' adherence to the MedDiet was classified as poor (\leq 3 points), medium (4–7 points) or high (\geq 8 points) according to their score (21,22).

Assessment of the outcome

We determined micronutrient intake adequacy for twenty micronutrients of known public health relevance, Zn; I; Se; Fe; Ca; K; P; Mg; Cr; Na; vitamin B₁; vitamin B₂; vitamin B₃; vitamin B₆; folic acid, vitamin B₁₂; vitamin C; vitamin A; vitamin D and vitamin E. To calculate the probability of intake adequacy, we compared the intakes of these nutrients with the estimated average requirements (EAR) when these were available or adequate intake levels, if not, as proposed by the Institute of Medicine⁽²³⁾. The traditional⁽²⁴⁾ and probabilistic⁽²⁵⁾ approaches were used. In the latter, the probability of adequacy for the usual intake of a nutrient was estimated from a z-score calculated as (derived nutrient intake - EAR)/SD of the EAR.

Evaluation of covariates

The baseline questionnaire collected information on sociodemographic and lifestyle variables, including physical activity and sedentary behaviour, as well as on personal and family medical records.

BMI was calculated using the ratio of reported weight (kg) to squared height (m²). The weight and height of the SENDO project participants reported by parents had been previously validated⁽²⁶⁾. Nutritional status was defined using sex- and age-specific BMI cut-off points based on International Obesity Task Force reference standards⁽²⁷⁾.





MedDiet and micronutrient inadequacy in child

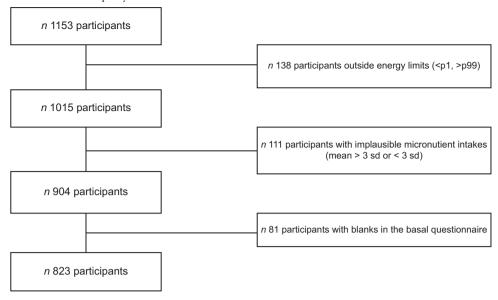


Fig. 1 Flow chart of participants

Physical activity was collected with a questionnaire that included seventeen activities and ten response categories, from never to ≥ 11 h/week. The Metabolic Task Equivalent (MET)-h/week for each activity was calculated by multiplying the number of MET of each activity by the weekly participation in that activity, weighted according to the number of months dedicated to each activity⁽²⁸⁾. Total physical activity was quantified by adding the MET-h/week of all the activities carried out during free time. Screen time was calculated as the average number of hours per day dedicated to watching TV, using a computer or playing video games. Parental knowledge of nutritional recommendations for children was assessed with questions about the recommended intake frequency of ten food groups (i.e. fruit, vegetables, dairy products). Parents had to choose among ten categories of response ranging from 'Never/ Almost never' to '≥ 6 or more times/day'. Each question was assigned 1 point if the answer complied with dietary recommendations and 0 points if not. The final score was expressed as a percentage, with a higher value meaning better knowledge about nutritional recommendations for children. For analysis, participants' knowledge was categorised as low (< 40%), moderate (40-70%) or high (> 70%). Parental attitudes towards their child's dietary habits were assessed with an 8-item questionnaire (i.e. I try to support my child to eat more fruit; I try to support my child to reduce the consumption of candies). Each question was assigned 1 point if the answer complied with dietary recommendations and 0 points if not. For analysis, parental attitudes were categorised as unhealthy (0-3 points), average (4-5 points) or healthy (6-8 points). Given the possibility of a change in diet due to the COIVD-19 pandemic, we divided the participants according to whether they completed the questionnaire before or after

the declaration of the state of alarm in Spain (14 March 2020).

Statistical analysis

Participants' socio-demographic characteristics were compared according to MedDiet adherence (low, medium or high). For descriptive purposes, we used means and standard deviations for quantitative variables and percentages for categorical ones. Nutritional characteristics of children's diet were also described and compared according to their adherence to MedDiet.

In the main analyses, we calculated the OR and 95 % CI for failing to meet the EAR of ≥ 3 micronutrients associated with the adherence to the MedDiet. The high adherence was used as the reference category. Crude and multivariate adjusted estimates were calculated through three progressively adjusted models. The first model was adjusted for sex (male or female), race (white v. others), nutritional status (underweight, normal weight, overweight/obese), total energy intake (quintiles), breast-feeding duration (no, < 6 months, 6-12 months, > 12 months) and pre- and postpandemic compliance, number of children and position held among siblings. The second model was adjusted for all the variables in model 1 plus maternal age (< 35 years, 35-40 years, > 40-45 years, > 45 years), maternal higher education (yes or no), parental knowledge about children's nutritional recommendations (low, medium or high score) and parental attitudes towards their child's dietary habits (unhealthy, average, healthy). Finally, the third model was adjusted for all the variables in models 1 and 2 plus physical activity (quintiles) and screen time (continuous).

Finally, we calculated the marginal effect of the adherence to the MedDiet, that is, the adjusted proportion



Table 1 Main characteristics of children in the SENDO project and their families (January 2015-June 2022). Numbers are mean (sp) or %

	Low adherence		Medium adherence		High adherence		
	n or Mean	% or sp	n or Mean	% or sp	<i>n</i> or Mean	% or sp	P _{for trend}
n	80	9.7 %	565	68.7%	178	21.6%	
Children's characteristics							
Sex (boys)	43	53.75	274	48.50	87	48.88	0.68
Age (years)	4.86	0.83	5.06	0.87	4.87	0.75	0.37
Race (White)	76	95.00	545	96.46	173	97.19	0.40
Screen time (h/d)	1.39	1.14	1.09	0.82	1.03	1.23	0.01
Physical activity (MET-h/d)	31.66	21.29	40.78	29.68	44.01	30.09	< 0.01
Birth weight (g)	3164-81	608.3	3240.06	548.5	3226.08	501.2	0.60
Breast-feeding duration							< 0.01
No breast-feeding	19	23.75	102	18.05	16	8.99	
< 6 months	33	41.25	159	28.14	45	25.28	
6–12 months	15	18.75	142	25.13	55	30.90	
> 12 months	13	16.25	162	28.67	62	34.83	
Z-score of BMI	0.11	1.09	0.05	1.17	0.04	1.10	0.68
Energy intake (kcal)	1720.84	447.2	2035.19	456·2	2180.51	458.5	< 0.01
Family characteristics	1720.04	777.2	2000-10	430·Z	2100.01	400.0	< 0.01
Mother's age (years)	39.45	4.32	39.97	4.40	39.95	4.27	0.52
Maternal higher education	58	72.50	460	81.42	151	84.83	0.02
Number of children	30	72.30	400	01.42	131	04.00	0.67
1–2	60	75.00	358	63.36	123	69-10	0.07
3	14	17·50	116	20.53	31	17.42	
<i>3</i> ≥ 4	6	7·50	91	20.53 16.11	24	13.48	
≥ 4 Position held among siblings	0	7.30	91	10.11	24	13.40	0.44
5 5	OF	31.25	207	36-64	66	37.08	0.44
The oldest or singleton	25						
2nd/3 or 2nd–3rd/4	7	8.75	97	17.17	22	12.36	
The youngest or ≥ 4th	48	60-00	261	46-19	90	50.56	0.00
Parental knowledge about dietary							0.02
recommendations for children							
Low (< 40 %)	24	30.00	132	23.36	30	16.85	
Medium (40–70 %)	49	61.25	364	64.42	113	63.48	
High (> 70 %)	7	8.75	69	12-21	35	19-66	
Parental attitudes towards							< 0.01
child's dietary habits							
Unhealthy (0-3 points)	13	16⋅25	24	4.25	6	3.37	
Average (4–5 points)	41	51.25	187	33.10	40	22.47	
Healthy (6–8 points)	26	32.50	354	62.65	132	74.16	

(and 95 % CI) of children with inadequate intake of ≥ 3 micronutrients in each category of adherence to the MedDiet.

To carry out a sensitivity analysis, calculations were repeated using the probabilistic method. A second sensitivity analysis was performed by adding to the micronutrient intake the content of the supplements reported by the participants.

Analyses were carried out using Stata 15.0 (Stata Corporation). All P-values are two-tailed. Statistical significance was established at the conventional cut-off point of P < 0.05.

Results

The main characteristics of participants and their parents are shown in Table 1. In this sample, 565 (68.7%) participants showed a medium adherence to the MedDiet, representing the largest group. Children with greater adherence to the MedDiet also presented overall healthier lifestyle indicators, such as less exposure to screens (P=0.01) and more time being physically active (P < 0.01), but also higher energy intake (P < 0.01). Similarly, longer breast-feeding duration was associated with greater adherence to the MedDiet (P < 0.01), which had previously been reported in this cohort⁽²⁹⁾. Parents who fed their children according to the MedDiet pattern showed greater knowledge about children's dietary recommendations (P=0.02) and displayed healthier attitudes towards their child's dietary habits (P < 0.01). Maternal education was marginally associated with higher adherence to MedDiet in this sample (P = 0.06).

Nutritional characteristics of the children's diet based on their adherence to the MedDiet are shown in Table 2. Children with the highest adherence to the MedDiet reported significantly higher (P < 0.001) consumption of carbohydrates, fibre, vegetables, fruit, legumes, cereals, potatoes, fish, nuts and eggs. On the contrary, those children presented significantly lower (P < 0.001) consumption of SFA and PUFA, fast food (P = 0.026) and other fats (P = 0.002).

We found significant associations between MedDiet adherence and energy-adjusted intake for seventeen out of





Table 2 Nutritional characteristics of children in the SENDO project (January 2015–June 2022) according to their level of adherence to the Mediterranean diet. Numbers are mean (SD) or %

	Low adherence		Medium adherence		High adherence		
	n or Mean	% or sp	n or Mean	% or sp	n or Mean	% or sp	P _{for trend}
n	80	9.7 %	565	68.7%	178	21.6%	
Total energy intake (kcal/d)	1720.84	447-2	2035-19	456-2	2180.51	458.5	< 0.001
Carbohydrate intake (% of TEI)	41.78	5.61	43.19	5.12	44.70	5.06	< 0.001
Protein intake (% of total energy)	17.24	2.17	17.12	2.20	16.82	1.98	0.079
Fat intake (% of total energy)	40.97	5.78	39.69	5.19	38.48	5.07	< 0.001
SFA intake (% of total energy)	12.00	2.24	11.29	2.12	10.48	1.98	< 0.001
MUFA intake (% of total energy)	15⋅19	3.99	15.31	3.50	15⋅36	3⋅19	0.734
PUFA intake (% of total energy)	5.39	1.84	4.62	1.03	4.55	0.92	< 0.001
SFA/MUFA intake	1.28	0.31	1.38	0.32	1.50	0.34	< 0.001
Fibre intake (g/d)	13.84	4.55	20.49	5.56	25.74	5.99	< 0.001
Food groups							
Vegetables (g/d)	84.83	52.0	186.7	102.4	255.6	98.8	< 0.001
Fruits (g/d)	201.5	186-3	357.7	196-2	500.6	219.8	< 0.001
Legumes (g/d)	21.64	15.04	32.36	18-35	36.04	17.64	< 0.001
Dairy (g/d)	435.6	249.7	490.3	237.5	482.5	233.4	0.326
Cereals (g/d)	59.54	30.27	75.63	38.30	88.44	39.12	< 0.001
Potatoes (g/d)	12.53	14.16	18.52	19.64	21.70	18.23	< 0.001
Meat (g/d)	126-22	41.56	134.74	45.53	131.17	47.96	0.778
Fish (g/d)	28.22	13.67	35.18	17-11	38.20	15.58	< 0.001
Nuts (g/d)	3.75	9.39	4.64	5.96	8⋅31	10⋅6	< 0.001
Bakery and sweets (g/d)	80.00	47.65	83.64	57.98	73.72	48.78	0.156
Sugar-sweetened beverages (g/d)	48.06	86.03	45.86	78-51	35.93	52.74	0.134
Fast Food (g/d)	58.95	28.62	60.26	26.61	54.85	32.25	0.098
Eggs (g/d)	16.74	9.26	19.64	10.25	21.35	7.78	0.001
Olive oil (g/d)	7.61	11.63	10.54	13.34	10.57	12.95	0.196
Other fats (g/d)	4.17	5.51	2.56	3.71	2.28	2.89	0.002

Table 3 Energy-adjusted micronutrient intake of children in the SENDO project (January 2015-June 2022) according to their level of adherence to the Mediterranean diet. Numbers are mean (SD)

	Low adherence		Medium ad	dherence	High adherence		
	n or Mean	% or sp	n or Mean	% or sp	n or Mean	% or sp	P _{for trend}
n	80	9.7 %	565	68.7%	178	21.6%	
Micronutrients							
Vitamin A (μg/d)	797.54	48.10	1045-43	17-69	1270.93	31.86	< 0.01
Vitamin C (mg/d)	78.62	6.26	135.44	2.30	180.54	4.15	< 0.01
Vitamin D (μg/d)	2.70	0.19	3.06	0.07	3.60	0.13	< 0.01
Vitamin E (mg/d)	8.96	0.31	8.08	0.11	8.97	0.20	0⋅15
Vitamin B ₁ (mg/d)	1.26	0.03	1.43	0.01	1.53	0.02	< 0.01
Vitamin B ₂ (mg/d)	1.83	0.05	2.07	0.02	2.07	0.04	< 0.01
Vitamin B ₃ (mg/d)	30.82	0.83	35.65	0.31	38.46	0.55	< 0.01
Vitamin B ₆ (mg/d)	1.89	0.05	2.27	0.02	2.55	0.03	< 0.01
Folic Acid (µg/d)	234.92	8.21	295.21	3.02	349.64	5.44	< 0.01
Vitamin B ₁₂ (mg/d)	4.32	0.15	4.69	0.05	4.79	0.10	0.02
Ca (mg/d)	1155.45	28.47	1166-86	10.47	1184.68	18.86	0.34
I (μg/d)	99.94	2.58	110.21	0.95	111.65	1.71	< 0.01
Fe (mg/d)	12.37	0.23	13.85	0.08	15.00	0.15	< 0.01
P (mg/d)	1481.65	72.55	1716.02	26.69	1902.35	48.06	< 0.01
Mg (mg/d)	258.18	4.77	296.08	1.75	328-91	3.16	< 0.01
Se (μg/d)	69.15	1.50	71.69	0.55	74.00	0.99	< 0.01
Zn (mg/d)	8.78	0.22	9.68	0.08	9.82	0.15	< 0.01
Cr (µg/d)	58.16	2.26	67.00	0.83	74.56	1.50	< 0.01
K (mg/d)	2828.53	64.08	3365.96	23.57	3793.19	42.44	< 0.01
Na (mg/d)	2902-25	96.13	2982-91	35.36	2921.13	63.68	0.84

the twenty micronutrients analysed, all except for vitamin E, Ca and Na. Of the micronutrients showing an association with MedDiet adherence, all showed a positive correlation (Table 3).

The spline at the top of Fig. 2 shows that, after adjusting for all the potential confounders, the change in micronutrient inadequacy (solid line) and the 95 % CI (dashed line) associated with the increase in MedDiet adherence





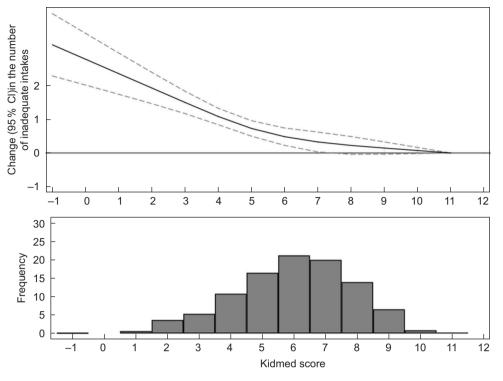


Fig. 2 The spline at the top shows that, after adjusting for all the potential confounders, the change in micronutrient inadequacy (solid line) and the 95 % CI (dashed line) associated with the increase in MedDiet adherence (assessing this variable as continuous) displayed a linear trend. MedDiet, Mediterranean Diet

 $\textbf{Table 4} \quad \text{OR and } 95 \% \text{ CI of failing to meet} \geq 3 \text{ micronutrients recommendations associated with adherence to the Mediterranean diet using the traditional method to define inadequate intake}$

	Low adherence		Mediur	m adherence	High adherence		
	OR	95 % CI	OR	95 % CI	OR	95 % CI	P _{for trend}
n	80		565		178		
%	9.7 %		68.7 %		21.6%		
Crude	21.30	9.36, 48.49	3.13	1.52, 6.45	1.00 (Ref.)		< 0.01
Multivariable adjusted model 1	12.74	4.76, 34.12	2.51	1.08, 5.86	1⋅00 (Ref.)		< 0.01
Multivariable adjusted model 2	10.89	3.73, 31.79	2.36	1.00, 5.53	1⋅00 (Ref.)		< 0.01
Multivariable adjusted model 3	9.85	3.33, 29.09	2.23	0.94, 5.29	1.00 (Ref.)		< 0.01

Model 1 is adjusted for sex (male or female), age (continuous), nutritional status (underweight, normal weight, overweight/obese), total energy intake (kcal), breast-feeding (no, < 6 months, < -12 months, > 12 months) and pre- and post-pandemic compliance, number of children and position held among siblings.

Model 2 is additionally adjusted for maternal age (< 35 years, 35–40 years, > 40–45 years, > 45 years), maternal higher education (yes or no), parental knowledge about child's nutritional recommendations (low, medium score or high) and parental attitudes towards child's dietary habits (unhealthy, average, healthy).

Model 3 is additionally adjusted for physical activity (tertiles) and screen time (tertiles).

(assessing this variable as continuous) displayed a linear trend. The regression analysis showed a slope of -0.26, meaning that, for every four extra points on the KIDMED index, there was a decrease of 1 micronutrient for which the intake was inadequate.

Table 4 represents the OR and 95 % CI for failing to meet the EAR for ≥ 3 micronutrients associated with the adherence to the MedDiet. Compared to participants with high adherence to the MedDiet, those with low and medium adherence showed 9·85-time (95 % CI 3·33, 29·09) and 2·23-time (95 % CI 0·94, 5·29), respectively, higher odds of having an inadequate intake of ≥ 3 micronutrients in the most adjusted model.

Figure 3 shows the marginal effect of the adherence to the MedDiet on the risk of failing to meet the EAR of ≥ 3 micronutrients. The adjusted proportion of children with ≥ 3 micronutrient inadequacies was significantly higher in the group of children with low adherence than in that with high adherence to the MedDiet. More specifically, the proportion of children who failed to meet the EAR of ≥ 3 micronutrient in the categories of low, medium and high adherence to the MedDiet was 27.3% (95% IC 19.3% to 35.3%), 13.5% (95% IC 11.1% to 15.9%) and 8.1% (95% IC 3.5% to 12.8%), respectively.

The robustness of these findings was assessed with two sensitivity analyses. The analysis that took into account the MedDiet and micronutrient inadequacy in child

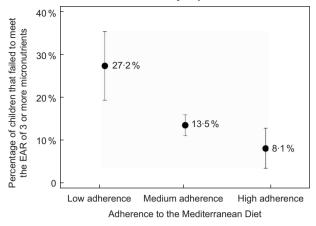


Fig. 3 Marginal effect of the adherence to the MedDiet on the risk of failing to meet the EAR of ≥ 3 micronutrients. MedDiet, Mediterranean Diet; EAR, estimated average requirement

content of the supplement intake reported by participants did not change the results (data not shown). On the other hand, the use of the probabilistic approach to define micronutrient inadequate intake resulted in higher estimates (see online supplementary material, Supplemental Table 1).

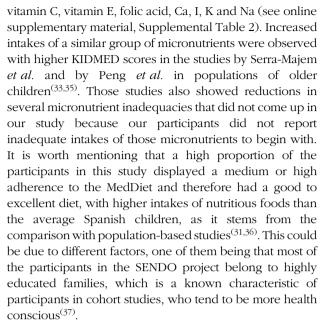
Discussion

This cross-sectional study based on 819 Spanish preschoolers examined the association between adherence to the MedDiet and micronutrient inadequacy. After adjusting for all the potential confounders, lower adherence to the MedDiet was directly associated with micronutrient inadequacy. Compared with children with high adherence to the MedDiet, those with medium or low adherence had 2·23 and 9·85-fold higher odds of failing to meet the EAR of 3 or more micronutrient, respectively.

To the best of our knowledge, this is the first study that focuses on the relation between the MedDiet and micronutrient inadequacy in young children. Our results are relevant from a public health perspective, given that a nonnegligible prevalence of micronutrient inadequacy persists among children and adolescents around the world⁽¹³⁾. This is also the case in more affluent regions such as Europe and North America, as recent studies have shown^(14,15,30–32).

These results are in line with previous findings, in both adults, adolescents and older children⁽¹⁶⁾. Moreover, as in the present study, a similar dose–response manner was found in previous ones^(33–35).

In our study, greater KIDMED scores were associated with higher intakes of vitamin A, vitamin C, vitamin D, vitamin B_1 , vitamin B_2 , vitamin B_3 , vitamin B_6 , vitamin B_{12} , folic acid, I, Fe, P, Mg, Se, Zn, Cr and K. Along with this, higher adherence to the MedDiet was associated with lower prevalence of inadequate intakes of vitamin A,



As previous studies have shown, the Mediterranean dietary pattern is of a high nutritional quality and displays an excellent micronutrient profile^(14,38,39). Thus, it is reasonable to think that the improvement in nutritional adequacy observed in this study could be due to the high content of nutrient-dense foods that make up the MedDiet. In fact, children with the highest adherence to the MedDiet showed higher consumption of vegetables, fruits, legumes, nuts, fish and eggs. They also presented lower consumption of sugar-sweetened beverages and bakery and sweets, which are foods known to be high in calories but low in nutrients.

Although it is true that we found a study population with a higher pattern of adherence to the MedDiet than the mean of other population-based studies in the paediatric population^(31,36), we consider that this is related to the fact that participants in cohort studies are usually health-conscious subjects with healthier lifestyles⁽³⁷⁾.

Overall, and despite its limitations, the KIDMED questionnaire appears to be useful to predict the risk of inadequate intake of micronutrients. The proportion of children with inadequate intakes decreased in a doseresponse manner as adherence to the MedDiet improved. More importantly, this proportion was notably low in the high adherence category.

This study is subject to certain limitations. First, the observational nature of our study does not enable us to eliminate possible residual confounding by unknown factors. Nonetheless, the study's large sample size and the fact that a substantial amount of information was collected from the participants have made it possible to adjust for many potential confounders. Moreover, the fact that our participants have parents with a high educational level reduces the possibility of confounding by socioeconomic factors⁽⁴⁰⁾ and improves the validity of the self-reported information. Second, since we used self-reported





information, the existence of measurement errors cannot totally be ruled out. A measurement error could lead to a misclassification bias in either the exposure (adherence to the MedDiet) or the outcome (micronutrient intake). Regarding the former, the possibility of misclassification was reduced by classifying the participants in three categories of exposure. Regarding the latter, the FFQ used in this study had been previously validated⁽⁴¹⁾, making measurement errors less likely. Additionally, since the participants did not know of the objective of the study, a potential misclassification bias would have been of the non-differential type, which would bias the results towards the null, making it more difficult to obtain statistically significant results. Fourth, our results refer to the probability of nutritional adequacy, not to actual nutrient deficiency, which can best be established through biomarkers of nutrient intake. Fifth, the sample of this study included only Spanish children, which may limit its external validity. However, we believe that the results could be generalised based on biological mechanisms and not based on statistical representativeness^(42,43). Lastly, due to the observational design of the study, the possibility of residual confounding by variables we did not account for (such as economic status) must be considered.

On the other hand, our study has several strengths. The large sample size and the large amount of information collected from the participants enabled a better control of confounding than some of the previous studies. Second, we excluded participants with energy or micronutrient intakes out of predefined ranges to avoid information bias. Third, we accounted for intra-cluster correlation between siblings in all the analyses, which is a common limitation of studies in paediatric populations. Fourth, the observed results are constant throughout the sensitivity analyses.

In conclusion, we found that lower adherence to the MedDiet was associated with a higher risk of nutritional inadequacy. Given the persistence of suboptimal micronutrient intake around the world, the Mediterranean dietary pattern may represent a promising option that could serve as a reference for Public Health nutrition policies to prevent micronutrient inadequacies in the paediatric population.

Acknowledgements

We thank all the participants of the SENDO project and their families for their invaluable collaboration with this project.

Financial support

This work did not receive any funding.

Conflict of interest

The authors declare no conflict of interest.

Authorship

All authors collected the data. A.O.O. and N.M.C. performed the statistical analyses. A.O.O. and E.F. wrote the first draft. All authors reviewed and commented on subsequent drafts of the manuscript.

Ethics of human subject participation

The SENDO project follows the rules of the Helsinki Declaration on Ethical Principles for Human Research, and its protocol was approved by the Ethical Committee for Clinical Research of Navarra (Pyto 2016/122). Participants' parents or legal guardians signed an informed consent before entering the study.

Supplementary material

For supplementary material accompanying this paper visit https://doi.org/10.1017/S1368980023002707

References

- 1. Fardet A & Rock E (2015) From a reductionist to a holistic approach in preventive nutrition to define new and more ethical paradigms. Healthc 3, 1054-63.
- Nestel PJ & Mori TA (2022) Dietary patterns, dietary nutrients and cardiovascular disease. Rev Cardiovasc Med 23, 17.
- 3. U.S. Department of Agriculture and U.S. Department of Health and Human Services (2020) Dietary Guidelines for Americans, 2020-2025. 9th ed. https://www.dietary guidelines.gov (accessed June 2023).
- 4. Vitale M, Masulli M, Calabrese I et al. (2018) Impact of a Mediterranean dietary pattern and its components on cardiovascular risk factors, glucose control, and body weight in people with type 2 diabetes: a real-life study. Nutrients 10,
- Guasch-Ferré M & Willett WC (2021) The Mediterranean diet and health: a comprehensive overview. J Intern Med 290, 549-566.
- Tosti V, Bertozzi B & Fontana L (2018) Health benefits of the Mediterranean diet: metabolic and molecular mechanisms. J Gerontol A Biol Sci Med Sci 73, 318–326.
- 7. D'innocenzo S, Biagi C & Lanari M (2019) Obesity and the Mediterranean diet: a review of evidence of the role and sustainability of the Mediterranean diet. Nutrients 11, 1306.
- 8. Martínez-González MA, Gea A & Ruiz-Canela M (2019) The Mediterranean diet and cardiovascular health. Circ Res 124, 779-798.
- Martinez-Gonzalez MA & Martin-Calvo N Mediterranean diet and life expectancy; beyond olive oil, fruits, and vegetables. Curr Opin Clin Nutr Metab Care 19, 401 - 407.
- World Health Organization Health Topics. Micronutrients. https://www.who.int/health-topics/micronutrients#tab=tab_1 (accessed June 2023).





- Panel E & Nda A (2013) Scientific Opinion on nutrient requirements and dietary intakes of infants and young children in the European Union. EFSA J 11, 1–103.
- 12. Ballesteros-Pomar MD & Arés-Luque A (2004) Déficit nutricionales carenciales (Nutritional deficiencies). *Endocrinol Nutr* **51**, 218–224.
- UNICEF (2019) State of the World's Children 2019: Children, Food and Nutrition 2019. https://www.unicef.org/media/ 63016/file/SOWC-2019.pdf (accessed September 2023).
- Hilger J, Goerig T, Weber P et al. (2015) Micronutrient intake in healthy toddlers: a multinational perspective. Nutrients 7, 6938–6955.
- Kaganov B, Caroli M, Mazur A et al. (2015) Suboptimal micronutrient intake among children in Europe. Nutrients 7, 3524–3535.
- Castro-Quezada I, Román-Viñas B & Serra-Majem L (2014)
 The Mediterranean diet and nutritional adequacy: a review. Nutrients 6, 231–248.
- Zazpe I, Santiago S, Romanos-Nanclares A et al. (2020) Validity and reproducibility of a semi-quantitative food frequency questionnaire in Spanish preschoolers — the SENDO project. Nutr Hosp 37, 672–684.
- Tuni OM, Carbajal Á, Vives CC et al. (2016) Tablas de Composición de Alimentos: Guía de Prácticas (Food Composition Tables: Practice Guide), 18th ed. Madrid: Piramide Ediciones Sa. Ciencia y técnica/Pirámide.
- Database (2010) TSFC Spanish Agency for Consumer Affairs, Food Safety and Nutrition. https://www.aesan.gob.es/en/ AECOSAN/web/home/aecosan_inicio.htm (accessed May 2023).
- Serra-Majem L, Ribas L, Ngo J et al. (2004) Food, youth and the Mediterranean diet in Spain. Development of KIDMED, Mediterranean Diet Quality Index in children and adolescents. Public Health Nutr 7, 931–935.
- Sahingoz SA & Sanlier N (2011) Compliance with Mediterranean Diet Quality Index (KIDMED) and nutrition knowledge levels in adolescents. A case study from Turkey. Appetite 57, 272–277.
- Rhee KE, DeLago CW, Arscott-Mills T et al. (2005) Factors associated with parental readiness to make changes for overweight children. Pediatrics 116, e94–101.
- Institute of Medicine (IOM) (2000) DRI Dietary Reference Intakes: Applications in Dietary Assessment. Washington, DC: National Academies Press (US). doi: 10.17226/9956.
- 24. Zazpe I, Sánchez-Taínta A, Santiago S et al. (2014) Association between dietary carbohydrate intake quality and micronutrient intake adequacy in a Mediterranean cohort: the SUN (Seguimiento Universidad de Navarra) Project. Br J Nutr 111, 2000–2009.
- Anderson GH, Peterson RD & Beaton GH (1982) Estimating nutrient deficiencies in a population from dietary records: the use of probability analyses. *Nutr Res* 2, 409–415.
- Oliver Olid A, Martín López L, Moreno Villares JM et al. (2021) Validation of the anthropometric data reported by parents of participants in the SENDO project. Nutr Hosp 38, 1162–1168.
- Cole TJ & Lobstein T (2012) Extended international (IOTF) body mass index cut-offs for thinness, overweight and obesity. *Pediatr Obes* 7, 284–294.

- Ainsworth BE, Haskell WL, Whitt MC et al. (2000) Compendium of physical activities: an update of activity codes and MET intensities. Med Sci Sports Exerc 32, 498–504.
- Oliver Olid A, Moreno-Galarraga L, Moreno-Villares JM et al. (2023) Breastfeeding is associated with higher adherence to the Mediterranean diet in a Spanish population of preschoolers: the SENDO project. Nutrients 15, 1278.
- United States Department of Agriculture ARS (2019) Usual Nutrient Intake from Food and Beverages. What We Eat in America, NHANES 2013–2016 Table A 1. https://www.ars. usda.gov/ARSUserFiles/80400530/pdf/usual/Usual_Intake_ gender_WWEIA_2013_2016.pdf (accessed June 2023).
- 31. Enalia E (2017) Estudio ENALIA 2012–2014: Encuesta Nacional de Consumo de Alimentos En Población Infantil y Adolescente (National Survey of Food Consumption in Children and Adolescents). https://www.aesan.gob.es/AECOSAN/docs/documentos/seguridad_alimentaria/gestio (accessed June 2023).
- Zaragoza-Jordana M, Closa-Monasterolo R, Luque V et al. (2018) Micronutrient intake adequacy in children from birth to 8 years. Data Childhood Obesity Project Clin Nutr 37, 630–637.
- Serra-Majem LI, Ribas L, García A et al. (2003) Nutrient adequacy and Mediterranean Diet in Spanish school children and adolescents. Eur J Clin Nutr 57, \$35–\$39.
- Ojeda-Rodríguez A, Zazpe I, Morell-Azanza L et al. (2018) Improved diet quality and nutrient adequacy in children and adolescents with abdominal obesity after a lifestyle intervention. Nutrients 10. 1500.
- Peng W, Berry EM & Goldsmith R (2019) Adherence to the Mediterranean diet was positively associated with micronutrient adequacy and negatively associated with dietary energy density among adolescents. J Hum Nutr Diet 32, 41–52.
- 36. Madrigal C, Soto-Méndez MJ, Hernández-Ruiz Á et al. (2022) Dietary intake, nutritional adequacy, and food sources of selected antioxidant minerals and vitamins; and their relationship with personal and family factors in Spanish children aged 1 to < 10 years: results from the EsNuPiStudy. Nutrients 14, 4132.
- Willet W (2012) Overview of Nutritional Epidemiology, 2nd ed. Oxford: Oxford University Press. pp. 9.
- Sánchez-Tainta A, Zazpe I, Bes-Rastrollo M et al. (2016) Nutritional adequacy according to carbohydrates and fat quality. Eur J Nutr 55, 93–106.
- Maillot M, Issa C, Vieux F et al. (2011) The shortest way to reach nutritional goals is to adopt Mediterranean food choices: evidence from computer-generated personalized diets. Am J Clin Nutr 94, 1127–1137.
- 40. Rothman KJ, Lash TL, VanderWeele TJ *et al.* (2020) *Modern Epidemiology*. Philadelphia: Wolters Kluwer.
- Zazpe I, Santiago S, de la O V et al. (2020) Validity and reproducibility of a semi-quantitative food frequency questionnaire in Spanish preschoolers — the SENDO project. Nutr Hosp 37, 672–684.
- Rothman KJ, Gallacher JEJ & Hatch EE (2013) Why representativeness should be avoided. Int J Epidemiol 4, 1012–1014.
- Rothman KJ (2014) Six persistent research misconceptions. J Gen Intern Med 29, 1060–1064.

