

Poor adherence to the Mediterranean diet is associated with increased likelihood of metabolic syndrome components in children: the Healthy Growth Study

Elena S George¹, Stavroula Gavrili², Catherine Itsiopoulos³, Yannis Manios^{4,5} and George Moschonis⁶,*

¹Institute for Physical Activity and Nutrition, School of Exercise and Nutrition Sciences, Deakin University, Geelong, Australia: ²Neonatal Intensive Care Unit, Alexandra University and State Maternity Hospital, Athens, Greece: ³School of Health and Biomedical Sciences, College of STEM, RMIT University, Melbourne, Australia: ⁴Department of Nutrition and Dietetics, School of Health Science and Education, Harokopio University, Athens, Greece: ⁵Institute of Agri-food and Life Sciences, Hellenic Mediterranean University Research Centre, Heraklion, Greece: ⁶Department of Dietetics, Nutrition and Sport, School of Allied Health, Human Services and Sport, La Trobe University, Melbourne, VIC 3086, Australia

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Abstract

Objective: To examine the associations between the level of adherence to the Mediterranean diet (MedDiet) with obesity, insulin resistance (IR), metabolic syndrome (MetS) and its components in schoolchildren.

Design: The Healthy Growth Study was a large epidemiological cross-sectional

Setting: School children who were enrolled in primary schools in four counties covering the northern, southern, western and central part of Greece were invited to

Participants: The study was conducted with a representative sample of 9-13-yearold schoolchildren (n 1972) with complete data. This study applied the KIDMed score to determine 'poor' (≤3), 'medium' (4-7) and 'high' (≥8) adherence of children to the MedDiet. The research hypothesis was examined using multivariate logistic regression models, controlling for potential confounders.

Results: The percentage of children with 'poor', 'medium' and 'high' adherence to the MedDiet was 64.8 %, 34.2 % and 1 %, respectively. Furthermore, the prevalence of obesity, IR and MetS was 11.6%, 28.8% and 3.4%, respectively. Logistic regression analyses revealed that 'poor' adherence to the MedDiet was associated with an increased likelihood for central obesity (OR 1.31; 95 % CI 1.01, 1.73), hypertriglyceridaemia (OR 2·80; 95 % CI 1·05, 7·46) and IR (OR 1·31; 95 % CI 1·05, 1·64), even after adjusting for several potential confounders.

Conclusions: The present study showed that approximately two-thirds of the examined sample of schoolchildren in Greece have 'poor' adherence to the MedDiet, which also increases the likelihood for central obesity, hypertriglyceridaemia and IR. Prospective studies are needed to confirm whether these are cause-effect associations.

Keywords Mediterranean diet children metabolic syndrome obesity insulin resistance

Globally, childhood obesity rates have increased dramatically over the past few decades⁽¹⁾. Epidemiological evidence has found childhood overweight and obesity to be a growing problem in Mediterranean countries, with the prevalence of obesity in the southern European region to be among the highest in Europe^(1,2). Greece, in particular, has been found to have the highest prevalence, of overweight and obesity, at approximately 20 %(2). It is well established that increased visceral fat mass levels and central adiposity present in obesity are associated with metabolic syndrome (MetS) which composes a cluster of risk factors including insulin resistance (IR), hypertension and dyslipidaemia⁽³⁾. Evidence also suggests that childhood obesity and related cardiometabolic risk factors are very

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likely to track into adulthood, thus increasing the risk for Type 2 Diabetes Mellitus and CVD⁽³⁾.

Unhealthy lifestyle habits, including physical inactivity, sedentary behaviour and poor diet quality, have been identified as the most important behavioural risk factors for obesity, MetS and its individual components⁽⁴⁾. Conversely, there is strong evidence surrounding the cardioprotective effect of the Mediterranean diet (MedDiet), with higher adherence demonstrating improved longevity and associations with reduced rates of atherosclerosis, MetS, CHD and inflammation in adults⁽⁵⁾. Studies assessing dietary intake within Mediterranean populations indicate that the traditional MedDiet is seldom being adhered to⁽⁶⁾. The change in dietary patterns, including straying from the traditional Mediterranean way of eating and its effect on health outcomes including obesity and related co-morbidities have not been well elucidated, especially in children. Furthermore, although adherence to the MedDiet has previously been reported to be poor in children in Greece⁽⁷⁾, the association of this poor adherence to the MedDiet with the MetS and its components has not been established.

The aim of the present study was to determine the adherence to the MedDiet using the KIDMed score, in 9-13-year-old schoolchildren in Greece, and to examine the association between the level of adherence to the MedDiet and obesity, IR, MetS and its components in this population.

Methods

Sampling

The Healthy Growth Study was a large-scale cross-sectional epidemiologic study initiated in May 2007 and completed in June 2009. The study population was composed of 9-13-year-old schoolchildren, attending the fifth and sixth grades of primary schools located in municipalities within the counties of Attica, Aitoloakarnania, Thessaloniki and Iraklio. The sampling of schools was random, multistage and stratified by parents' educational level and total population of students attending schools within municipalities of these counties (based on Census data), as described in more detail elsewhere (8). The study population was regionally representative of the 9-13-year-old school children living in the four counties under study. Nonetheless, these counties are scattered throughout the Greek territory, covering the northern (i.e. Thessaloniki), central (i.e. Attica), western (i.e. Aitoloakarnania) and southern (i.e. Iraklio-Crete) parts of Greece. This combined with the random, multistage and stratified sampling procedures followed to recruit our sample are indicative of the representativeness of our population. The sampling procedure yielded seventy-seven primary schools, representative of the total number of schools in the counties under study, which responded positively when they were invited to participate in the study. An extended letter explaining the aims of the study and a

consent form for taking full measurements were provided to all parents or guardians having a child in these schools. Signed parental consent forms were collected for 2656 children (response rate 64·1%), whose parents allowed their participation to the study. Assessment between responders and non-responders based on initial screening data indicated that there were no significant differences between groups for children's weight and parental education level (data not shown).

Dietary intake

Dietary intake data were obtained by trained dietitians and nutritionists via morning interviews with the children at the school site, using a twenty-five item FFQ and three 24-h recalls. More specifically, children's habitual dietary intake was assessed with an FFQ, which included food items from all food groups, as well as discretionary foods that are usually consumed by children in this age group. The 24-h recalls were collected through face-to-face interviews from all study participants. Children were asked to describe the type and amount of foods and beverages consumed, during the previous 24 h for two consecutive weekdays and one weekend day. The main prerequisite for the 24-h recalls was that the previous day was a usual day according to the participant's perception. To improve the accuracy of food description, standard household measures (cups, tablespoons, etc.) and food models were used to define amounts. At the end of each interview, the interviewers, who were dietitians rigorously trained to minimise the interviewer's effect, reviewed the collected data with the respondent in order to clarify entries, servings and possible forgotten foods. Food intake data were analysed using the Nutritionist V diet analysis software (version 2.1, 1999, First Databank), which was extensively amended to include traditional Greek recipes⁽⁹⁾. The database was updated with nutritional information for foods available in Greece including processed foods provided by independent research institutes, food companies and fast-food chains.

Physical activity

Physical activity was assessed using a standardised questionnaire validated with accelerometers by a subsample of the Healthy Growth study⁽¹⁰⁾. In this study, children completed the questionnaire for two consecutive weekdays and one weekend day. Reported activities were assessed as moderate-to-vigorous physical activity provided that they were of intensity higher than four metabolic equivalents and included activities such as bicycling, rhythmicgymnastics, dancing, basketball, soccer, athletics, tennis, swimming, jumping rope and general participation in active outdoor games. Given the age group, moderateto-vigorous physical activity was defined as continuous physical activities causing sweating and heavy breathing for periods longer than 15 min, but with occasional breaks



in intensity, rather than the strict aerobic definition of twenty continuous minutes appropriate for adults⁽¹¹⁾.

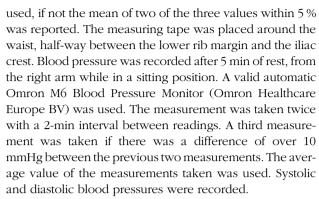
KIDMed score

Children's adherence to the MedDiet was assessed using the KIDMed score (see online Supplementary Table 1)⁽¹²⁾, via a combination of data stemming from the FFQ and the 24-h recalls. The score combined sixteen questions that summarise the characteristics of the Mediterranean dietary pattern. A score of +1 was used for 'yes' responses to positive questions, and a score of -1 was given to 'ves' responses to negative questions; 0 was given to 'no' responses. Total possible scores ranged between -4 and 12. The data derived from the FFO were used for the calculation of the sub-scores of those KIDMed components related to children's habitual consumption of fruits and fruit juice, vegetables, fish, fast food, legumes, pasta or rice, breakfast cereals or grains, nuts, olive oil, yogurt and/or cheese, sweets and candy. Considering that the information derived from the FFO was adequate to cover thirteen out of sixteen components for the calculation of the KIDMed score, the information for the remaining three components resulted from the data stemming from the three 24-h recalls. In this regard, the 24-h recall data related to skipping breakfast and consumption of dairy products and baked goods or pastries for breakfast were extracted from the 24-h recalls.

Total KIDMed scores equal or lower than 3 were indicative of 'poor' adherence to the MedDiet, scores between 4 and 7 reflected 'medium' adherence, while scores higher than 8 were indicating 'high' adherence⁽¹²⁾. For the purpose of this study and to ensure a more meaningful analysis, moderate and high adherence were merged as there were a very small number of children with high adherence.

Anthropometry and physical examination

Participants underwent a physical examination by two trained members of the research team. The protocol and equipment used were the same in all schools. Weight was measured to the nearest 10 g using a Seca digital scale (Seca Alpha, Model 770). Students were weighed without shoes in the minimum clothing possible. Height was measured to the nearest 0.1 cm using a commercial stadiometer (Leicester Height Measure, Invicta Plastics) with the participant standing barefoot, keeping shoulders in a relaxed position, arms hanging freely and head in Frankfurt horizontal plane. Weight and height were used to calculate BMI using Quetelet's equation (weight (kg)/height (m)²). The International Obesity Task Force cut-off points (13,14) were used to categorise participants as 'underweight', 'normal weight', 'overweight' or 'obese'. Waist circumference (WC) was measured in triplicate to the nearest 0.1 cm with the use of a non-elastic tape (Hoechstmass) with the student standing, at the end of a gentle expiration. If the three measurements were within 5 % of each other the mean was



Furthermore, one well-trained and experienced paediatrician in each prefecture determined pubertal maturation (Tanner stage) after thorough visual inspection of breast development in girls and genital development in boys⁽¹⁵⁾.

Biochemical indices

Blood samples were obtained for biochemical and haematological screening tests between 08-30 and 10-30 after a 12-h overnight fast. Reminders were distributed the previous day to both parents and children to ensure compliance with fasting. Professional staff performed venipuncture to obtain a maximum of 23 ml blood. Part of the blood was collected in test tubes with no added anticoagulant, where it was allowed to clot for approximately 2 h, as this was designated for serum separation. Clotted blood was centrifuged at 3000 rpm for 15 min, and the collected serum was divided into aliquots and stored at -80 °C. All serum samples were transported in dry ice to the Laboratory of Nutrition and Clinical Dietetics at Harokopio University, where central storage of back-up samples at -80 °C took place.

Total cholesterol, HDL-cholesterol and TAG were determined in duplicate using commercially available enzymatic colorimetric assays (Roche Diagnostics SA) on an automated analyser (Roche/Hitachi Modular). LDL-cholesterol was calculated by the Friedewald equation⁽¹⁶⁾.

Plasma glucose was determined using commercially available enzymatic colorimetric assays (Roche Diagnostics SA). Serum insulin was determined by a Chemiluminescence immunoassay (Kyowa Medex Ltd, for Siemens Diagnostics USA). IR was measured through homoeostasis model assessment (HOMA-IR) $^{(17)}$. This index was calculated using fasting glucose and fasting insulin as follows: HOMA-IR = (fasting insulin (µunits/ml) x fasting glucose (mmol/l)/22·5. HOMA-IR > 3·16 $^{(18)}$ was used as a cut-off point to define insulin-resistant schoolchildren. HOMA-IR has been validated as a surrogate measure of IR in non-diabetic children, with studies showing correlations as high as 0·91 with the hyperinsuline-mic-euglycemic clamp, a frequently sampled intravenous glucose tolerance test $^{(19)}$.

MetS was defined using the criteria proposed by the International Diabetes Federation for children and adolescents. Children with MetS had abdominal obesity





 $(WC \ge 90^{th} \text{ percentile by age and sex})$ and the presence of two or more of the following clinical features: elevated TAG $(\geq 1.7 \text{ mmol/L})$; low HDL-cholesterol (< 1.03 mmol/L); high blood pressure (systolic blood pressure ≥ 130 mm Hg and/ or diastolic blood pressure ≥ 85 mm Hg and/or diagnosis of hypertension) and elevated plasma glucose (≥ 5.6 mmol/L and/or diagnosis of type 2 diabetes)⁽²⁰⁾.

Socio-economic and other information collected from the parents

Data on the socio-economic background of the families having at least one child participating in the study were collected from the parents (most preferably from the mother) during scheduled face-to-face interviews at school. For those parents not able to attend the meetings (approximately 5 % of the total sample), data were collected via telephone interviews. All interviews were conducted by research team members who were rigorously trained to minimise interviewer's effect by using a standardised questionnaire. More specifically, the data collected by primary caregivers included demographic characteristics such as parent's age and nationality; socio-economic characteristics such as educational level (years of education) and mean annual family income over the last 3 years (€/year). Lifestyle characteristics such as smoking habits and anthropometric characteristics such as mother's height and weight were also collected.

Statistical analysis

Continuous variables were summarised as mean (SD), while categorical variables were presented as frequencies (n) and percentages (%). The Kolmogorov-Smirnov test was used to examine the normality of the distribution of all continuous variables. Differences of mean values between groups were examined using Students' t-test (or the non-parametric Mann-Whitney test in the case of non-normally distributed continuous variables), while associations between categorical variables were examined by using the χ^2 test and the twosample z-test for proportions whenever appropriate. Logistic regression analyses were performed to evaluate the associations between KIDMed score with MetS, MetS components, IR and obesity in children. More specifically, two different models were applied: model 1 unadjusted and model 2 - adjusted for children's sex, Tanner stage, urbanisation degree (i.e. urban, semiurban or rural), maternal educational level, physical activity and dietary daily energy intake. A post hoc power calculation based on available sample size showed that there was more than 80 % statistical power to detect effect sizes in the multivariable logistic regression models. The results of logistic regression models were presented as OR and 95 % CI. The level of statistical significance was set at P < 0.05, while all reported P-values were two-tailed. All statistical analyses were conducted using the Statistical Package for Social Sciences (SPSS) software for Windows (version 24.0, SPSS Inc).

Results

Table 1 presents the main descriptive characteristics of the study population. The total number of children with full data, included in the current statistical analysis, was 1972, with approximately 50% of children being girls. The mean age of the study participants was $11 \cdot 1 (0.6)$ years, with the vast majority of children (65.2%) living in urban areas. Regarding sex differences, dietary energy intake and physical activity levels were significantly higher in boys compared with girls (P < 0.001). In addition, BMI, WC, serum LDL-cholesterol and plasma glucose levels were also higher in boys compared with girls (P < 0.05), while girls had higher levels of serum TAG, serum insulin, HOMA-IR and diastolic blood pressure compared with boys (P < 0.001). A similar pattern was also observed for the level of biological maturation, since the majority of girls were categorised at higher Tanner stages compared with boys (P < 0.001). When comparisons between poor and medium and high adherence to MedDiet were carried out, the results indicated that there was a higher energy intake with better compliance to MedDiet (P < 0.001), while higher adherers to the MedDiet were more physically active (P < 0.001). They also had significantly lower BMI and WC and more favourable lipid profiles (Table 1).

Figure 1 depicts the percentages of children, in the total sample and by sex, with 'poor', 'medium' and 'high' adherence to the MedDiet based on the KIDMED score. According to these percentages, 64.8 %, 34.3 % and 1 % of children were found to have 'poor', 'medium' and 'high' adherence to the MedDiet, respectively. As there was such a low frequency of 'high' adherence to the MedDiet medium and high adherence groups were pooled together for analyses. Relevant percentages, as in the total sample, were also observed for boys and girls, with no statistically significant differences between sexes.

Table 2 summarises the prevalence of the MetS and its individual components for the total sample, by sex and by adherence to the MedDiet. The overall prevalence of the MetS was 3.4%, while regarding its individual components, 15.8% of children were found to have central obesity, 1.7%hypertriglyceridaemia, 7.3 % low HDL-cholesterol concentrations, 17.7% hyperglycaemia, 24.8% systolic hypertension and 7.4% diastolic hypertension. In addition, 28.8% and 11.6% of children were insulin resistant and obese, respectively. Regarding sex differences, a higher percentage of girls compared with boys had low HDL-cholesterol concentrations (8.8 % v. 5.9 %, P = 0.013), diastolic hypertension $(8.9\% \ v. \ 5.9\%, \ P=0.011)$ and IR $(35.1\% \ v. \ 22.5\%,$ P < 0.001). On the contrary, a higher percentage of boys were hyperglycaemic (22.7% v. 12.6%, P<0.001) and





Table 1 Descriptive characteristics of the examined population (total sample and by sex)

	Total	sample	e (n 19	72)	Boys (<i>n</i> 990)				Girls (<i>n</i> 982)				KIDMED score ≤ 3 (Poor Adherence) (n 1277)				KIDMED score > 3 (Medium/High Adherence) (<i>n</i> 695)					
	Mean	SD	n	%	Mean	SD	n	%	Mean	SD	n	%	Mean	SD	n	%	Mean	SD	n	%	P-value† (sex comparison)	P-value† (MedDiet adherence comparison)
Socio-demographics																						
Children's age (Years)	11.1	0.6			11.1	0.7			11.1	0.6			11.2	0.7			11.1	0.6			0.884	0.604
Maternal education (Years)	12.6	3.3			12.8	3.2			12.4	3.3			12.3	3.3			13.1	3.1			0.018	<0.001
Urbanisation Level Rural Semi-urban			391 296	19·8 15·0			188 142				203 154				771 196				_	4 74·0 0 14·4	0.371	<0.001
Urban Behavioural				65.2				66.7				63.6				24.3				1 11.7		
Energy intake (kcal/	1799-1	552.2			1903-0	582.0			1694.4	499-2			1726.4	523.6			1932.7	578-2			<0.001	<0.001
d) MVPA (min/day) <i>Anthropometrics</i>	65.6	61.1			76.5	64.4			55.7	55.4			62.5	60-4			71.4	61.9			<0.001†	<0.001†
Body weight (kg)	45.3	11.0			45.7	11.2			45.0	10.9			45.6	11.4			44.8	10.3			0.160	0.140
Height (cm)	148.7	7.8			148-4	7.3			149.1	8.2			148.5	7.9			149.1	7.7			0.057	0.105
BMľ (kg/m²)	20.3	3.8			20.6	3.9			20.1	3.7			20.5	4.0			20.0	3.5			0.004	0.009
Waist circumfer-	69.0	9.7			70.3	10.1			67.7	9⋅1			69.5	10.2			68∙1	8.8			<0.001	0.003
ence (cm) Biochemical																						
Serum TAG (mg/dl)	62.8	28.4			59.4				66.3	29.0			64.2	30.1			60.3	24.7			<0.001	0.003
Serum total choles- terol (mg/dl)	166.7	32.0			168.0				165.5	32.3			166.7	31.7			166-8	32.6			0.083	0.983
Serum HDL-choles- terol (mg/dl)	59.3	15.0			59.9	15.2			58-6	14.8			58-1	14.3			61.6	16.0			0.060	<0.001
Serum LDL-choles- terol (mg/dl)	94.9	26.7			96.2	27.1			93.6	26.3			95.8	26.8			93.1	26.5			0.030	0.032
Plasma glucose (mg/dl)	92.6	9.9			91.4	9.4			90.3	8.6			91.3	9.3			91.7	9.6			<0.001	0.401
Serum insulin (mU/mL)	12.2	8.4			11.1	8.8			13.4	7.7			12.5	8.2			11.7	8.7			<0.001	0.030
HOMA-IR Clinical	2.7	1.9			2.5	2.1			2.9	1.7			2.8	1.8			2.6	2.2			<0.001	0.074
Systolic blood pres-	121.1	13.2			121.3	13.0			121.0	13.4			121.2	13.5			121.0	12.8			0.604	0.692
sure (mmHg) Diastolic blood pressure (mmHg)	70.2	9.9			69.4	9.8			71.1	9.8			70-6	10.2			69.6	9.3			<0.001	0.044
Tanner stage																						
Stage 1			664	33.7			438	44.2			226	23.0			436	34.1			22	8 32.8	<0.001	0.775
Stage 2			805	40.8			439				366				522				28	3 40.7		
Stage 3			355	18.0			97	9.8			258	26.3			228	17.9				7 18.3		
Stage 4			122	6.2			15	1.5			107	10.9			77	6.0			4			
Stage 5			26	1.3			1	0.1			25	2.5			14	1.1			1:	2 1.7		

^{*}P-values were derived from Student's *t*-test in the case of normally distributed continues variables and the χ^2 test in the case of categorical variables. †P-value was derived from the non-parametric Mann–Whitney test. MVPA = -moderate-to-vigorous intensity physical activity; HOMA-IR = homoeostatic model assessment insulin resistance.

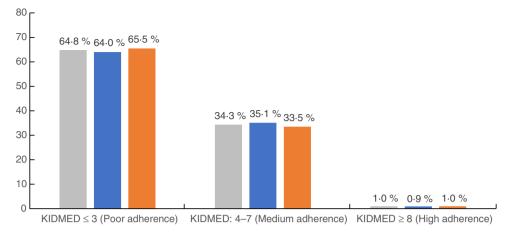


Fig. 1 (colour online) Percentages of children with 'poor', 'medium' and 'high' adherence to the Mediterranean diet, based on the KIDMED score in the total sample and by sex. ■, Total sample; ■, boys; ■, girls

obese (13.6 % v. 9.5 %, P = 0.004) compared with girls. Regarding differences between children with different adherence to the MedDiet, the percentages of children with central obesity (17.5 % v. 12.5 %, P = 0.003), hypertriglyceridaemia (2.2 % v. 0.7 %, P = 0.015) and IR (30.8 % v. 25.1 %,P = 0.008) were higher in those with poor compared with those with medium or high adherence. Furthermore, children with medium/high adherence to the MedDiet had a higher maternal educational level (13.0 (sd 3.1) v.12.4 (sd 3.3)years, P < 0.001), energy (1932.7 (sd 578.1) v. 1726.4 (sd 523.6) kcal/day, P < 0.001)and time spent doing moderate-to-vigorous physical activities $(71.4 (sd 61.9) \ v. 62.5 (sd 60.4) \ min/day, \ P = 0.002)$, compared with children with poor adherence to the MedDiet. In addition, a higher percentage of children with medium/high adherence to the MedDiet were living in urban areas, compared with their peers with poor adherence to the MedDiet (74.0 % v. 60.4 %, P < 0.05). On the contrary, a lower percentage of children with medium/high adherence to the MedDiet were living in rural areas, compared with children with poor adherence to the MedDiet (11.7% v. 24.3%, P < 0.05) (data not shown in Tables).

Table 3 summarises the results from the two logistic regression models that were performed to examine the associations between KIDMed score with MetS, MetS components, IR and obesity in the examined population of children. According to the results derived from model 1 (unadjusted model), 'poor' adherence to the MedDiet was found to be associated with an increased likelihood for central obesity (OR 1·49; 95 % CI 1·14, 1·94), hypertrigly-ceridaemia (OR 3·09; 95 % CI 1·19, 8·05) and IR (OR 1·49; 95 % CI 1·33, 1·64). The significant associations observed in model 1 were also retained in model 2, after adjusting for sex, Tanner stage, physical activity levels and total dietary energy intake, family urbanisation degree and maternal educational level. More specifically in model 2 (adjusted

model), 'poor' adherence to the MedDiet was found to increase the likelihood for central obesity, hypertriglyceridaemia and IR by 1.31~(95~%~CI~1.01,~1.73), 2.80~(95~%~CI~1.05,~7.46) and 1.31~(95~%~CI~1.05,~1.64) times, respectively.

Discussion

The aim of the present study was to determine adherence to the MedDiet using the KIDMed score, in 9–13-year-old school children in Greece, and to evaluate the association between MedDiet adherence and obesity, IR, MetS and its individual components. Based on the findings from this study, adherence to MedDiet in this population was poor and was associated with an increased likelihood of central obesity, hypertriglyceridaemia and IR.

In this study, we focused on paediatric populations where there are currently little data assessing the effect of a MedDiet and how this impacts metabolic outcomes⁽²¹⁾. An increased prevalence of obesity, and subsequently MetS in paediatric populations⁽¹⁾, is known to result in elevated future risk of CVD. Therefore, it is important that modifiable risk factors such as diet are explored to determine if the incidence and outcomes related to MetS can be mediated to reduce the risk of CVD. Subsequently, this may help to reduce related morbidity and mortality in adulthood. Whilst there is extensive literature to support the efficacy of the MedDiet in adults for the prevention and management of MetS⁽²²⁾, the evidence is scarce in paediatric populations. The MedDiet has been shown to efficiently decrease occurrence of MetS(23), specifically in children and adolescents and an inverse association between adherence to the MedDiet and obesity has also been reported⁽²⁴⁾. Given the extensive protective role of MedDiet extended upon from the results from this study, further investigation of the MedDiet within children is warranted.







Table 2 Prevalence of metabolic syndrome (MetS), MetS components, insulin resistance and obesity in the total sample, by sex and by adherence to the mediterranean diet

		sample 972)	Boys	(n 990)	Girls	(n 982)	scoi (poor er	MED re ≤ 3 adher- ace) 277)	scor (mediu adhe	MED re > 3 um/high rence) 695)		
	n	%	n	%	n	%	n	%	n	%	P-value† (sex comparison)	P-value† (MedDiet adherence comparison)
Metabolic Syndrome WC > 90 th percentile $T_g \ge 150$ mg/dl (or ≥ 1.7 mmol/L) HDL-cholesterol < 40 mg/dl (or <1.03 mmol/L) FPG ≥ 100 mg/dl (or ≥ 5.6 mmol/L) Hypertension based on SBP Hypertension based on DBP Insulin resistance (HOMA-IR > 3.16) Obesity (BMI \ge 95th percentile)	68 311 33 144 349 489 145 565 228	3.4 15.8 1.7 7.3 17.7 24.8 7.4 28.8 11.6	32 162 14 58 225 245 58 222 135	3.2 16.4 1.4 5.9 22.7 24.7 5.9 22.5 13.6	36 149 19 86 124 244 87 343 93	3.7 15.2 1.9 8.8 12.6 24.8 8.9 35.1 9.5	50 224 28 95 225 318 104 391 159	3.9 17.5 2.2 7.4 17.6 24.9 8.1 30.8 12.5	18 87 5 49 124 171 41 174 69	2.6 12.5 0.7 7.1 17.8 24.6 5.9 25.1 9.9	0.598 0.468 0.367 0.013 <0.001 0.959 0.011 <0.001 0.004	0.060 0.003 0.015 0.751 0.902 0.884 0.068 0.008

WC, waist circumference; To, TAG; FPG, fasting plasma glucose; SBP, systolic blood pressure; DBP, diastolic blood pressure; HOMA-IR, homoeostatic model assessment insulin resistance. †P-values were derived from the χ^2 test in the case of categorical variables.

Table 3 Likelihood of metabolic syndrome (MetS), increased levels of MetS components, insulin resistance and obesity in Greek children (n 1972) with poor adherence to the Mediterranean diet pattern (KIDMED score ≤ 3)*

	Metabolic Syndrome		WC > 90th percentile		Tg ≥ 150 mg/dl (≥1·7 mmol/L)		HDL-choles- terol < 40 mg/dl (< 1.03 mmol/L)		FPG ≥ 100 mg/dl (≥5·6 mmol/L)		Hypertension based on SBP		Hypertension based on DBP		Insulin resistance (HOMA-IR > 3.16)		Obesity (BMI ≥ 95th percentile)	
	OR	95 % CI	OR	95 % CI	OR	95 % CI	OR	95 % CI	OR	95 % CI	OR	95 % CI	OR	95 % CI	OR	95 % CI	OR	95 % CI
Model 1 (Unadjusted) KIDMED score																		
>3 (Medium/High Adherence)	1.00		1.00		1.00		1.00		1.00		1.00		1.00		1.00		1.00	
≤3 (Poor Adherence) Model 2 (Adjusted†) KIDMED score	1.61	0.98, 2.64	1.49	1.14, 1.94	3.09	1.19, 8.05	1.06	0.74, 1.52	0.99	0.77, 1.25	1.04	0.84, 1.29	1.41	0.97, 2.06	1.33	1.08, 1.64	1.29	0.96, 1.74
>3 (Medium/High Adherence)	1.00		1.00		1.00		1.00		1.00		1.00		1.00		1.00		1.00	
≤3 (Poor Adherence)	1.49	0.85, 2.61	1.31	1.01, 1.73	2.80	1.05, 7.46	0.99	0.69, 1.45	1.16	0.90, 1.51	1.01	0.81, 1.27	1.26	0.86, 1.86	1.31	1.05, 1.64	1.16	0.85, 1.59

WC, waist circumference; T_g , TAG; FPG, fasting plasma glucose; SBP, systolic blood pressure; DBP, diastolic blood pressure; HOMA-IR, homoeostatic model assessment insulin resistance. *Cells in bold indicate statistically significant OR.

[†]Adjusted for sex and Tanner Stage (stages 1-5), maternal educational level (years), urbanisation level (rural, semi urban and rural), time spent to - - (min/d) and total dietary daily energy intake (kcal).



To determine the level of children's adherence to the MedDiet, the present study used the KIDMed score, which is the most widely used index in studies conducted with children and adolescents(21). The results from this study indicated that approximately two-thirds of children had 'poor' adherence to the MedDiet, while only 1 % of children had 'high' adherence. These results demonstrate that despite Greece being located within the Mediterranean basin, dietary intake has become Westernised, a notion that is in line with other literature reporting low adherence to the MedDiet in Greek children^(21,25,26). When comparing these results to other paediatric cohorts in the Mediterranean region, the levels of adherence to the MedDiet were lower than those reported in a study for Cypriot children, where only 6.7 % of children had 'high' adherence to the MedDiet, while 37 % were found to have 'poor' adherence⁽²⁷⁾ A systematic literature review on children and adolescents, which included thirty-eight studies that used KIDMED score to assess adherence to the MedDiet⁽²¹⁾, indicated that adherence to the MedDiet varied across Mediterranean countries⁽⁷⁾ More specifically, the review indicated that across the whole population, 'poor' adherence was lowest in Spanish children $(1.6\%)^{(28)}$ and highest in Greek adolescents $(62.8\%)^{(29)}$, 'moderate' adherence was highest in rural Italian adolescents (73.8%)(30) and lowest in Greek adolescents (28.0 %)(31), while 'high' adherence was lowest in Greek adolescents (4.3%)⁽⁷⁾ and highest in Spanish children⁽³²⁾. These findings confirm the results reported by the present study, highlighting that approximately two-thirds of children in Greece have a 'poor' adherence to the MedDiet.

Large epidemiological data from the 1999 to 2002 NHANES database, a US representative sample within a comparable demographic, indicated a prevalence of MetS in paediatric populations ranging from 2 to 9 % depending on the diagnostic criteria used⁽³³⁾. The rates of MetS reported within the present study are not dissimilar to that of the NHANES, despite the difference in locality with 3.4 % of children are diagnosed with the MetS, thus falling within the range reported in a seemingly Westernised obesogenic population. Data on MetS in previous studies have demonstrated a higher prevalence in boys compared with girls, although there was no significant difference between sexes from the results herein⁽³⁴⁾. However, what was alarming was a large number of children, despite not meeting the criteria for the MetS who had established MetS components. Rates of central adiposity and IR were particularly high (almost 16% and 30%, respectively) in the total sample, and obesity and hyperglycaemia were significantly higher in boys. Furthermore, our findings showed that IR was higher in girls compared with boys, which was expected considering the earlier biological maturation (reflected by higher Tanner Stage scores) observed in girls, which occurs during this life stage of preadolescence (35). The prevalence of IR observed in the present study is notably higher compared with the rates reported in a systematic literature review, which showed that IR (based on HOMA-IR) in children ranged from 2 to 20 %^(17,36). These findings are concerning given that IR and subsequent hyperinsulineamia have been proposed as the underlying mechanisms driving MetS, as well as other diseases which relate to obesity and/or the MetS, including polycystic ovary syndrome in females, non-alcoholic fatty liver disease, obstructive sleep apnoea and certain types of cancer⁽³⁷⁾.

There was an overall low prevalence of MetS (3.4%) in this paediatric population which makes sense given that MetS is not an actual clinical condition in children and adolescents, thus the focus by paediatric health professionals is mainly given to the individual components of MetS (i.e. central obesity, dyslipidaemia, hyperglycaemia and hypertension). In fact that there is no consensus for the definition of the MetS in paediatric populations to date. In this cohort of children, however, there was poor adherence to the MedDiet and high rates of MetS components. The rates of unfavourable cardiometabolic risk factors were higher in children with poor adherence to the MedDiet compared with children with 'moderate' and 'high' adherence. The MedDiet is characterised by a high intake of plant-based foods featuring vegetables, fruits, wholegrains, legumes, nuts, seeds and extra virgin olive oil. Smaller amounts of fish, white meat and fermented dairy are also included with red meat and sweets consumed least often. This high fibre diet is low in saturated fat and high in monounsaturated fat, predominantly attributable to the extra virgin olive oil, which is the main culinary fat (38). It is hypothesised that the phytonutrient rich, plant-based foods and high amounts of unsaturated fats included in this diet help mediate blood lipids, glycaemic control, while the anti-inflammatory and antioxidant properties of the MedDiet reduce inflammation and oxidation that also contribute to the MetS^(38,39). This has been demonstrated extensively in adult populations (40), but the present study is one of few studies reporting this potential relationship in children. In this study, poor adherence to the MedDiet was associated with central adiposity and thus we hypothesise that the MedDiet may be satiating⁽⁴¹⁾. This has previously been recognised due to the palatability (38) and high healthy fat composition, and as such, a higher level of adherence to the diet may regulate energy balance and thus regulate weight status.

Although there is some evidence examining the relationship between childhood obesity with adherence to the MedDiet, there are limited studies reporting associations between adherence to the MedDiet with central adiposity, IR and components of the MetS. A systematic review assessing children and adolescents included ten studies that reported significant negative associations between the adherence to the MedDiet and BMI or obesity status⁽²¹⁾. Only two of these studies provided evidence of a negative association between MedDiet adherence and WC in children, thus confirming the findings observed in the present study^(42,43). However, most of the relevant literature includes observational studies, while only two prospective

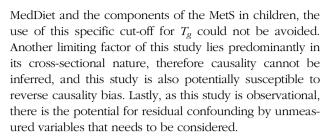




studies examined associations between MedDiet adherence with BMI or WC in children, but they did not report any significant findings. The only known randomised intervention study was conducted in Mexico⁽⁴⁴⁾ and reported a positive effect of the MedDiet on children's BMI. Insulin resistance was also assessed in this trial, and their results indicated that MedDiet was effective at reducing fasting glucose⁽⁴⁵⁾. Children in this study also improved their TGA levels⁽⁴⁶⁾ which is consistent with the negative association between adherence to the MedDiet and hypertriglyceridaemia observed in our study findings. In another intervention study in a paediatric population after 12 months, total and LDL-cholesterol levels improved, but there was no change in the serum concentrations of TGA⁽⁴⁷⁾. Therefore, additional prospective and intervention studies are required to examine the potential cardioprotective effect of the MedDiet on the cardiometabolic profile in children and adolescents, to determine if there is consistency with the positive relationships observed in adults as part of intervention studies.

Strengths and limitations

The strengths of this study lie in the large epidemiological data set provided from the Healthy Growth Study, a representative sample of children from four counties within the wider region of Greece. However, there are a number of limitations associated with the KIDMed score, these include that the score was inspired by instruments developed for adults and there are no specific guidelines for the Mediterranean Diet in non-adult populations. Furthermore, while some analysis has shown that the KIDMed score is positively associated with key nutrients (Ca and vitamin C⁽²⁶⁾), additional assessment of the validity of the score with biological outcomes is warranted. KIDMed score was calculated by combining data that were collected using information extracted on FFQ and 3 x 24 h recalls (2 week days and one week end), this represents a limitation of this study posing potential self-reporting bias, although it worth highlighting that the KIDMed score has been calculated this way in many other studies⁽²¹⁾. In addition, the FFQ and food recalls are based on self-reported dietary intake and pose a potential cause of study bias. However the FFQ has not been validated but was designed to assess children's habitual intake of the main and most popular foods and beverages consumed in this age group, while it has been repeatedly used in the past in other published work from the Healthy Growth Study, indicating reasonable and strong associations between children's food habits and clinical and biochemical markers of nutritional status (46-48). One additional limitation of our study also stems from the use of a high cut-off for defining hypertriglyceridaemia ($T_g > 150 \text{ mg/dl or} \ge 1.7$ mmol/L), since this may have resulted in an overestimation of the OR in the association of hypertriglyceridaemia with the KIDMED score. However, as the aim of the present study was to examine the associations between adherence to the



In conclusion, the present study was one of the very few to investigate adherence to the MedDiet and associations with obesity, IR, MetS and its components in a paediatric population. The results showed that approximately two-thirds of the examined population of Greek schoolchildren have 'poor' adherence to the MedDiet, which also leads to a higher likelihood for central obesity, hypertriglyceridaemia and IR. Additional prospective or intervention trials are needed to infer whether there is causal association between MedDiet and MetS components. If this relationship is confirmed, identifying strategies to increase adherence to the MedDiet among children in Greece may be an effective approach to tackle the cardiometabolic complications reflected in the components of the MetS in this population.

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Supplementary material

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