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RESEARCH ARTICLE

The assessment of Chronic Obstructive Pulmonary Disease (COPD) and its relation to the Mediterranean diet in Moroccan adults - BOLD study

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Authorship

AB conceived the study idea, its design, led the analyses and interpretation of the data, and drafted the manuscript. KE contributed to the conception of the study, and contributed to the review and the editing. IH contributed to the proofreading review and the editing. MMSD and ZH contributed to the conception of the study. ME, CN and MCB contributed to the study design and to the data collection. KE contributed to the study design and to the data collection. Conceived the study idea, its design, and led the analyses and interpretation of the data and supervised the drafting. All authors have read and approved the manuscript.

Abstract

Objective: The protective effect of the Mediterranean diet (MD) is known for several diseases, but the evidence in low- and middle-income countries was still missing. This article assesses the impact of MD and its components on Chronic Obstructive Pulmonary Disease (COPD) among Moroccan adults.

Methods: in population-based cross-sectional study, A total of 744 adults with acceptable spirometry according to the GOLD guidelines were randomly selected from a sample frame, of Moroccan adults lived in the areas of the Saïis district-Fez city. Dietary data were collected through a validated food frequency questionnaire. Mediterranean Diet Score (MDS) was used to assess Adherence to the Mediterranean food model, A value of 0 or 1 has been assigned to each of the eight indicated food components according to their beneficial or deleterious effect on health.

Results: Although no significant associations were found between COPD and the overall MD. score, associations were found between some of the MD components and COPD when stratifying for overall MDS adherence level (low, middle, high adherence). For the high adherence group, the high consumption of cereals, fruits and nuts were inversely associated with COPD risk with OR = 0.64; 95% CI = 0.26-0.89, and OR= 0.67; 95% CI = 0.44-0.96, respectively. The high consumption of meats and dairy products was positively associated with the risk of COPD, with OR = 1.37; 95% CI = 1.22-2.87, and OR= 1.83; 95% CI = 1.21-2.76, respectively.

Conclusion: The results of this study confirmed previous results showing significant associations of COPD risk with some components of MDS. Extensive studies are needed to explore MDS components better and suggest more effective interventions to maintain healthy eating habits and reduce COPD risk.

Introduction

Morocco is a North African country located on the southern shore of the Mediterranean Sea with nearly 34 million inhabitants¹. This country suffers from the consequences of a deviation from the Mediterranean Diet (MD), and it has experienced a nutritional transition over the past two decades, presented by a shift in its dietary pattern², which is associated with an increased burden of non-communicable diseases³. Several nutritional and behavioral risk factors such as unhealthy diet, smoking, and irregular physical activity have been suggested as causing illness and lung injuries such as asthma and chronic obstructive pulmonary disease (COPD). According to the world health organization (WHO)⁴, COPD is a lung disease presented by chronic obstruction of lung airflow that interferes with normal breathing and is not fully reversible. COPD is among the leading causes of morbidity and mortality worldwide³. Its prevalence in Morocco reaches 12.6% of people over 40⁵. It is the fourth leading cause of death and the fifth leading cause of hospitalization in men⁶. Smoking is the major risk factor for COPD, while MD and its components may play an essential role in preventing this disease⁷.

MD consists of a mixture of foods with beneficial or deleterious effects on lung function⁸. Adherence to the MD is described as high consumption of beneficial components, including vegetables, legumes, whole grains, nuts, fruits, olive oil, moderate consumption of dairy products, eggs, poultry, wine, and low consumption of red meat⁹. These beneficial components have anti-inflammatory and antioxidant properties and play an essential role in maintaining the balance between beneficial/deleterious, oxidant/antioxidant, and inflammatory/anti-inflammatory properties^{10,11}.

Evidence from epidemiological data shows that MD could be a protective factor against pulmonary diseases through its beneficial components, such as phytochemicals with antioxidant properties. As such, MD improves respiratory functions and ameliorates the quality of life of COPD patients^{8,12}.

Most studies described the association between intake of nutrients or isolated foods and COPD risk. However, it does not appear easy to discern the contribution of one food or one nutrient to the well-being of individuals. Hence, the importance of evaluating the diet as a whole, considering the function of different components (foods and nutrients) cannot be dissociated from the rest of the daily diet. However, complementary approaches providing evidence of the possible modulating effect of dietary patterns on lung health and the

protective or deleterious effect of individual dietary factors on the risk of COPD are limited.

In the light of the literature, the relation between nutritional status or eating habits and COPD in Morocco remains uncertain. Therefore, a complete analysis assessing the impact of MD compliance and its components on COPD risk could shed light on the potential of an MD in the battle against the increasing COPD epidemic.

Methods

Study design and Participants

Among adults aged ≥ 40 years, a BOLD cross-sectional population survey was carried out in Fez city, Morocco¹³. The design and rationale of the BOLD survey methodology have been published elsewhere¹⁴. Participants were randomly selected, from a sample frame, of Moroccan adults living in the Saïis district. Adults were eligible if they were ≥ 40 years old and lived in a randomly selected household. A total of 760 adults with acceptable spirometry values were included in this study.

Data Collection

The study data was collected through the BOLD questionnaire, developed from pre-existing validated questionnaires and used in international surveys administered by trained staff. The questionnaire includes socio-demographic data: age, gender, and educational level (collected in years and divided into two classes: < 6 years of school or ≥ 6 years of school). In addition, data about some potential risk factors were also collected: smoking (classified into three classes: non-smokers, ex-smokers, and current smokers); Body Mass Index (BMI, kg/m²) as a quantitative variable, and the Waist Circumference (WC), which was measured with soft tape at the Middle of the iliac and umbilicus. In addition, abdominal obesity was assessed using the sex-specific WC cut-off points. Normal WC was defined by $WC \leq 102$ cm in men ≤ 88 cm in Women, and abdominal obesity ($WC > 102$ cm in men and > 88 cm in women)¹⁵.

Spirometry

Pulmonary function measurements (FEV₁ and FVC) were performed by trained and certified technicians using an NDD easyone spirometer¹⁶. FEV₁ (the amount of Forced expiratory volume in one second) refers to the maximum amount of air that the subject can forcibly expire during the first second following maximal inhalation. FVC (Forced vital capacity) refers to the maximum amount of air that can be exhaled when blowing out as fast as

possible. FEV1/FVC ratio was used in diagnosing COPD according to the GOLD guidelines¹⁷. COPD stage 1 or higher was defined as post-bronchodilator Forced Expiratory Volume in 1 second to Forced Vital Capacity (FEV1/FVC) ratio less than 70%. Each spirogram was examined and scored using guidelines from the American Thoracic Society (ATS) and European respiratory society (ERS)¹⁶. The measurement quality was checked by the Burden of Obstructive Lung Disease (BOLD) Pulmonary Function Reading Center at Imperial College London, UK. Only 760 adults were selected due to conforming to spirometry quality.

Dietary intake assessment

Trained interviewers collected dietary information using a validated Food Frequency Questionnaire (FFQ) that includes 32 food sections and 255 traditional and modern foods commonly consumed in Morocco¹⁸. Standard units or reference servings were specified for each food. The questionnaire included eight categories of intake frequency, ranging from "never or less than once a month" to "four or more times a day." First, each food intake frequency was converted into an average daily intake for each participant. Next, the information was converted into a daily consumption frequency of each food using a Moroccan food composition table (MFCT)¹⁹ and local household units.

Calculation of the Mediterranean diet score

Adherence to the Mediterranean food model was assessed by calculating the Mediterranean Diet Score (MDS)⁹. The MDS included eight food components (Cereals, Olive Oil, Fruits and Nuts, Legumes, Vegetables, Fish, Meat, and Dairy Product). The alcohol consumption component was removed because it was not applicable due to the cultural context. A value of 0 or 1 has been assigned to each of the eight indicated components according to their beneficial or deleterious effect on health. The gender-specific median of each component was used as a threshold. For foods with a beneficial effect on health, such as vegetables, legumes, olive oil, fruits and nuts, cereals, and fish, a value of 1 is assigned for consumption higher than the median; and for foods with a deleterious effect, such as dairy products and meat, a value of 1 was assigned for consumption lower than the median. The scores for each food component were then added together to obtain a total MDS ranging from 0 (minimum MDS) to 8 (maximum MDS). This MDS was divided into 3 MD-adherence (MDA) categories: Low-MDA (MDS ranged from 1 to 3), Moderate-MDA (MDS 4-5), and High-MDA (MDS 6-8).

Statistical analyses

Descriptive statistics, such as means and frequencies, were performed to describe all variables. Continuous variables were shown as Means \pm Standard Deviation (SD), and categorical variables as percentages. Chi-square and Student t-test were used adequately to assess the association between COPD and each of these variables.

Bivariate analysis was used to assess the association between the average weekly servings for the eight MDS components (independent variables) and the COPD status (dependent variable) in the whole population and separately in the three MDA categories. In addition, differences among COPD and non-COPD subjects were evaluated using a Student's t-test.

Crude and adjusted multivariate regression models were used to analyze the association between COPD and the MD as a categorical variable (the high MDA category as a reference group) and as a quantitative variable MDS. These analyses were run for the whole population and also stratified by sex (men adjusted for tobacco (smokers population) while women were not adjusted for tobacco because they are predominantly non-smokers (99%)).

On the other hand, the binary logistic regression models were used to estimate the association between COPD risk and the MD components in the whole population and in each of the three MDA-categories. All adjusted multivariate analyses considered the following potential confounders: age, gender, smoking habits, abdominal obesity, and total energy intake. The low and high estimates of TEI values were excluded from the database (1% at each end of the distribution) to control for total energy intake (TEI) as a potential confounding factor.²⁰

All analyzes were performed with SPSS version 21 (SPSS Inc., Chicago, IL, USA). *P*-values lower than 0.05 were considered statistically significant.

Results

In this cross-sectional study, the average age of the 744 participants was 55.2 (\pm 10.20) years, and there were slightly more women (53.2%) than men (46.8 %). The prevalence of smoking was approximately 9.5% for current smokers (21.0% in men and 0.5% in women) and 17.2% for ex-smokers (36.7% in men and 0.5% in women). The average MDS was 4.38 (\pm 1.21), it was 4.36 (\pm 1.22) in men, and 4.40 (\pm 1.20) in women. Adherence to the MD was mainly moderate to high (37.9% and 39.0%, respectively). The overall prevalence of COPD was 14.1%. It is more

prevalent in men (19.2%) than women (9.7%). COPD was more prevalent in the group of subjects with moderate adherence to MD (18.8%) than those with high and Low adherence to MD (12.4% and

9.3%, respectively). The main demographic, clinical, and nutritional characteristics according to the MDA categories are shown in Table 1.

Table 1: Demographics. Clinical and nutritional characteristics according to the three categories of Mediterranean diet adherence (low, moderate, and high); BOLD study. Moroccan Center (N=744)

	Low adherence (MD Score 1-3) N= 172		Moderate adherence (MD Score 4-5) N=282		High Adherence (MD Score 6-8) N=290		P*	P**
Age (Years) (M±SD)	53.98	±9.80	56.33	±11.02	54.36	±9.86	NS	0.03
Gender								
Men n (%)	81	47.1%	116	41.1%	147	50.7%	NS	0.01
Women n (%)	91	52.9%	166	58.9%	143	49.3%		
Educational level (Years)								
<6 years	132	76.7%	208	73.8%	220	75.9%	NS	NS
>6 years	40	23.3%	74	36.2%	63	24.1%		
Smoking status								
Current-smokers n (%)	16	9.3%	19	6.7%	29	10.0%		
Ex-smokers n (%)	27	15.7%	44	15.6%	70	24.1%	0.02	<0.01
Non-smokers n (%)	129	75.0%	219	77.7%	191	65.9%		
BMI (Kg/m ²) (M±SD)	28.19	±5.66	28.20	±5.05	27.41	±5.17	NS	NS
Abdominal-Obesity (WC) n (%)	101	58.7%	160	56.7%	132	45.5%	0.01	<0.001
TEI (Kcal/d)	3496.17	±1042.11	3614.56	±1056.49	3878.12	±1115.92	<0.001	<0.001
COPD n (%)								
FEV1 post (M±SD)	2.66	±0.74	2.50	±0.73	2.68	±0.70	0.004	0.002
FVC post (M±SD)	3.36	±0.88	3.22	±0.87	3.43	±0.85	NS	0.01
FEV1 post/FVC post (M±SD)	79.15	±6.68	77.31	±8.78	78.34	±8.62	NS	NS

P *: P-value for low compared to high adherence to M.D.; P **: P-value for moderate compared to high adherence to MD: with. Student t-test to compare means for continuous variables; Chi-Square Test for categorical data

TEI: Total Energy Intake in kilocalories per day. BMI: Body Mass Index in kilograms by meters squared.; COPD: Chronic Obstructive Pulmonary Disease; MD: Mediterranean Diet; W.C.: Waist Circumference; FVC: Forced Vital Capacity. FEV1 post: a spirometry post-bronchodilator forced expiratory volume in one second.

In our population, when compared to high MDA, participants with moderate MDA were mainly women, non-smokers, low total energy intake (TEI), high abdominal obesity (WC), low FEV1 and FVC, respectively, and higher prevalence of COPD. In addition, when compared to high-MDA, low-MDA was marked by a high rate of abdominal obesity (WC), a low average of TEI, low FEV1, and low COPD prevalence.

Table 2 presents the mean servings per week for the different food components of the MDS and the mean of TEI according to the three MDA-categories of people with or without COPD. The population's

COPD status was marked by a low average serving of cereals, vegetables, fruits and nuts, legumes, and TEI, whereas COPD status was linked to a high average serving of dairy products. The average servings of cereals, vegetables, fruits, and nuts were higher in non-COPD subjects compared to COPD ones in the high as well as the moderate MDA categories. Among low and high-MDA compared to non-COPD subjects, the average servings of dairy products were lower in COPD participants, whereas the average servings of meat were higher in COPD subjects with high-MDA. An increased average TEI was observed in COPD subjects in the moderate-MDA group only.

Table 2. Average weekly servings for the different MDS components and Total Energy Intake in people with or without COPD living in Fez. BOLD study (N = 744)

	All			Low adherence (MD Score 1-3)			Moderate adherence (MD Score 4-5)		High adherence (MD Score 6-8)			
	COPD-	COPD +	p	COPD-	COPD +	p	COPD-	COPD +	P	COPD-	COPD +	p
	639	105		N=165	N=16		N=229	N=53		N=254	N=36	
MDS-components												
Mean (±SD)												
Cereals	21.6±6.08	20.85±5.84	<0.001	20.07±7.59	18.29±8.60	<0.001	21.64±4.94	20.12±5.33	0.04	23.04±5.33	20.99±5.12	0.01
Olive oil	11.30±5.57	11.31±5.89	N.S.	9.86±6.55	10.76±7.31	N.S.	12.14±6.15	11.68±5.76	N.S.	11.13±5.43	10.9±5.45	NS
Vegetables	8.33±3.95	7.81±3.16	<0.001	9.17±3.54	10.31±3.53	NS	7.77±3.85	6.74±3.12	0.01	8.19±4.34	7.95±3.24	0.05
Fruits and nuts	6.65±4.64	5.91±4.2	<0.001	5.95±3.61	6.53±2.24	NS	6.31±3.40	5.49±3.34	0.02	7.13±5.97	6.02±5.98	0.02
Legumes	1.19±1.21	0.94±0.82	<0.001	0.85±0.66	0.7±0.70	NS	0.88±0.72	0.83±0.67	N.S.	2.59±2.34	2.39±1.04	NS
Fish	1.78±1.38	1.95±1.11	NS	1.41±0.99	1.46±0.95	N.S.	1.74±1.07	1.8±1.01	N.S.	1.99±1.75	2.2±1.39	NS
Meat	4.87±2.35	4.77±2.31	N.S.	5.45±3.90	6.74±3.12	N.S.	4.84±1.92	4.86±2.30	NS	3.95±2.17	4.74±2.06	0.03
Dairy products	7.08±5.31	8.22±5.43	<0.01	7.96±7.79	9.3±5.42	0.04	8.78±5.83	8.9±5.77	NS	4.74±4.62	7.36±5.19	0.004
TEI	3744.33±1102.79	3473.5±982.46	<0.001	3522.03±1035.13	3279.29±1053.02	NS	3671.09±1068.86	3345.56±824.35	0.02	3872.39±1057.09	3664.5±1261.80	NS

COPD: Chronic Obstructive Pulmonary Disease; MDS Mediterranean diet Score; SD Standard Deviation; TEI: Total Energy Intake; NS: Not Significant

Associations between the MDA categories and COPD risk are presented in Table 3. No significant associations were found between these MDA categories and COPD risk in the population or both genders. However, significant associations were found between specific components of the MDS and COPD in the whole population and each of the MDA categories (Table 4). Overall, and regardless of age, sex, smoking habits, abdominal obesity as well

as total energy intake, there was a statistically significant inverse association between the individual components of the MDS and COPD risk for cereals, fruits and nuts, and vegetables, while a positive association with dairy-products was observed. In addition, the risk of COPD decreased in men for olive oil and legumes and increased for meat. In contrast, these components were not associated with COPD risk in women

Table. 3: Crude and Adjusted models for MDA categories and COPD risk by gender (N=744)

MDA categories	Model 1	Model 2
	OR (95% CI)	OR (95% CI)
Overall (N=744)		
High MDA* (N=290)	1	1
Moderate MDA (N=282)	1.04 (0.59 ; 1.83)	0.93 (0.51 ; 1.70)
Low MDA (N= 172)	1.22 (0.72 ; 2.08)	1.31 (0.75 ; 2.27)
Men (N=344)		
High MDA (N=147)	1	1
Moderate MDA (N=116)	1.01 (0.49; 2.05)	0.75 (0.35 ; 1.62)
Low MDA (N=81)	1.25 (0.65 ; 2.40)	1.18 (0.60 ; 2.34)
Women (N=400)		
High MDA (N=143)	1	1
Moderate MDA (N=166)	1.43 (0.52 ; 3.90)	1.49 (0.52 ; 4.28)
Low MDA (N=91)	1.52 (0.58 ; 3.99)	1.71 (0.62 ; 4.73)

*Mediterranean Diet Adherence

Model 1: brut odds ratio

Model 2: overall M.D. adjusted for age, gender, smoking habits, abdominal obesity, and total energy intake.

Model 2: In men, adjusted for age, smoking habits, abdominal obesity, and total energy intake (all smokers were men).

Model 2: In women, adjusted for age, abdominal obesity, and total energy intake.

In the low-MDA group, COPD risk decreased with increased cereals consumption. However, this association disappeared when stratified by gender. Moderate-MDA was marked by the inverse relationship between the COPD risk and the consumption of cereals, fruits, nuts, and vegetables, while a positive association was observed with meat consumption. This association remains negative for legumes and vegetables in men but positive for

meat and dairy products. Finally, in the high-MDA group, COPD risk was inversely associated with cereals, fruits, and nuts while positively associated with consuming meat and dairy products. By contrast, only olive oil and legumes were inversely associated with COPD risk in men, while dairy products were positively associated with COPD risk in women.

Table 4: Logistic binary regression models of M.D. food categories and their association with COPD in 3 levels of adherence, for the total population and by gender (N=744)

	Whole population	CI 95%	Low Adherence	CI 95%	Moderate Adherence	CI 95%	High Adherence	CI 95%
All^a								
Cereals	0,58	(0,38; 0,89)*	0,57	(0,38 ; 0,87)*	0,66	(0,41 ; 0,98)*	0,64	(0,26 ;0,89)*
Olive Oil	1,45	(0,60 ; 3,46)	0,63	(0,27 ; 1,50)	1,08	(0,43 ; 2,71)	0,84	(0,06 ; 10,54)
Fruits and Nuts	0,56	(0,36 ; 0,86)*	1,01	(0,98 ; 1,04)	0,92	(0,91 ; 0,99)*	0,67	(0,44 ; 0,96)*
Legumes	0,96	(0,63 ; 1,46)	0,94	(0,59 ; 1,49)	1,03	(0,66 ; 1,60)	1,13	(0,31 ; 4,04)
Vegetables	0,84	(0,26 ; 0,87)*	1,19	(0,77 ; 1,86)	0,97	(0,96 ; 0,99)*	0,65	(0,07 ; 5,72)
Fish	1,41	(0,66 ; 1,54)	1,86	(0,91 ; 3,82)	1,01	(0,66 ; 1,54)	2,00	(0,55 ; 7,22)
Meat	1,34	(0,88 ; 3,04)	0,97	(0,64 ; 1,47)	2,10	(1,00 ; 4,50)*	1,37	(1,22 ; 2,87)*
Dairy Products	1,26	(1,08 ; 1,89)*	3,77	(0,99 ; 14,24)	0,86	(0,45 ; 1,64)	1,83	(1,21 ; 2,76)*
For Men^a								
Cereals	0,73	(0,39 ; 1,37)	0,64	(0,21 ; 1,94)	0,62	(0,23 ; 1,66)	0,40	(0,20 ; 2,75)
Olive Oil	0,48	(0,24 ; 0,95)*	0,72	(0,07 ; 7,94)	0,44	(0,08 ; 2,59)	0,12	(0,06 ; 0,59)*
Fruits and Nuts	1,09	(0,07 ; 17,86)	1,17	(0,51 ; 2,24)	0,52	(0,21 ; 1,31)	0,41	(0,04 ; 5,39)
Legumes	0,42	(0,34 ; 0,94)*	0,66	(0,29 ; 2,14)	0,34	(0,12 ; 0,96)*	0,13	(0,01 ; 0,62)*
Vegetables	0,79	(0,14 ; 4,33)	1,15	(0,39 ; 3,43)	0,33	(0,14 ; 0,79)*	0,82	(0,57 ; 11,97)
Fish	1,39	(0,93 ; 1,88)	1,03	(0,33 ; 3,22)	1,88	(0,77 ; 4,58)	1,84	(0,27 ; 12,85)
Meat	1,73	(1,24 ; 6,19)*	0,87	(0,32 ; 2,35)	2,30	(1,15 ; 5,45)*	2,40	(0,39 ; 8,57)
Dairy Products	1,67	(1,15 ; 5,43)	1,75	(0,59 ; 5,24)	1,49	(1,19 ; 4,71)*	1,26	(0,22 ; 7,38)
For Women^a								
Cereals	0,66	(0,53 ; 1,75)	0,51	(0,10 ; 2,58)	0,51	(0,18 ; 1,48)	0,87	(0,67 ; 2,03)
Olive Oil	0,82	(0,69 ; 5,32)	0,68	(0,32 ; 1,78)	1,08	(0,23 ; 5,82)	1,04	(0,27 ; 6,95)
Fruits and Nuts	0,99	(0,97 ; 1,00)	0,87	(0,24 ; 3,18)	0,57	(0,20 ; 1,62)	0,71	(0,44 ; 1,16)
Legumes	0,92	(0,46 ; 4,15)	0,77	(0,22 ; 2,73)	1,83	(0,63 ; 5,33)	1,23	(0,62 ; 4,24)
Vegetables	0,99	(0,97 ; 1,00)	1,15	(0,28 ; 4,75)	0,89	(0,31 ; 2,59)	0,81	(0,53 ; 3,68)
Fish	1,53	(0,72 ; 3,47)	1,07	(0,31 ; 3,80)	1,75	(0,62 ; 4,96)	2,16	(0,23 ; 6,42)
Meat	1,07	(0,29 ; 5,03)	0,46	(0,14 ; 1,55)	1,25	(0,43 ; 3,64)	1,06	(0,56 ; 5,56)
Dairy Products	1,12	(0,71 ; 5,78)	2,94	(0,71 ; 12,26)	0,66	(0,22 ; 2,03)	1,75	(1,31 ; 3,46)*

^aRegression analysis Adjusted for age, sex, smoking habits, abdominal obesity, and total energy intake; *Regression analysis Adjusted for age, smoking habits, abdominal obesity, as well as total energy intake; *Regression analysis Adjusted for age, abdominal obesity, as well as total energy intake.

Discussion

This cross-sectional study was designed to examine the relationship between COPD, the Mediterranean diet, and its components in Moroccan adults living in Fez. The key findings indicated that, regardless of the MDA category, the risk of COPD decreased with the increased consumption of cereals. The same association was found for fruits and nuts in the whole population, as well as the moderate and high MDA categories, while for vegetables in the whole population and the moderate MDA category only. Interestingly, the COPD risk decreased with the increasing consumption of olive oil and legumes in men only. On the other hand, the COPD risk increased with the increasing consumption of meat in moderate and low-MDA categories and the high consumption of dairy products in the whole population and the high-MDA category.

According to scientific and epidemiological literature, the high MDS reflects high intakes of vegetables, fruit and nuts, whole grains, polyunsaturated fatty acids (PUFA),

monounsaturated fatty acids (MUFA), and low intakes of red and processed meat, refined grains, empty calories and sodium^{21,22}. Individual consumption of some of these foods has been linked to better lung function, while others have been shown to have a deleterious effect on respiratory function⁷. Several results of previous prospective and retrospective studies^{8,12,23} support the hypothesis that greater adherence to the Mediterranean diet and its beneficial components may blunt the adverse respiratory effects and the risk of developing COPD.

In our study, cereals intake was negatively related to COPD status regardless of MDA categories, and this result can be explained by the fact that cereals constitute the main food in almost all Moroccan dishes. In addition, being at the bottom of the MD-Pyramid²¹, cereals are the primary source of nutritional energy, protein, iron, vitamins, and fiber²⁴. Their potential beneficial effects on human health are numerous: they lower blood cholesterol and glucose levels, reduce toxins and preserve intestinal

flora's beneficial balance²⁵. Also, their effect on pulmonary functions has been proven by clinical and epidemiological tests as protective factors against smoking effects and repair damaged lung tissue^{26,27}. Our study was in line with many others: A prospective study exploring the role of dietary fiber intake and the risk of COPD in American women and men concluded that the consumption of fiber, particularly cereal fiber, was negatively associated with the risk of newly diagnosed COPD²⁷. In addition, the MORGEN study²⁸ suggests a beneficial independent effect of whole grains (> 45g per day) on COPD.

Another finding from our study showed that higher consumption of vegetables, fruit, and nuts, both the main components of the MDS, is associated with improved lung function for moderate and high-MDA. The same results have been confirmed by previous epidemiological literature, according to which the consumption of antioxidants in fresh fruits and vegetables could compensate for the damage caused by tobacco and inhaled particles, could delay the premature aging of the lung, reduce the production of mucosa and therefore also reduce cough^{10,29}. Indeed, several epidemiological studies have found that a "prudent" diet, rich in fruits and vegetables, has a beneficial effect on respiratory function, especially FEV1, and is linked to a low prevalence of newly diagnosed COPD^{30,31}. Furthermore, other studies have concluded that daily intake of vegetables and fruits can slow the decline in respiratory function even among former smokers^{10,31}. For example, a Swedish prospective cohort exploring the effect of fruit and vegetable consumption on the incidence of COPD concluded that each daily serving of fruit and vegetables reduced the risk of COPD by 8% and 4%, respectively, among current and ex-smokers³². Another beneficial effect of consuming fruits and vegetables was related to weight gain and obesity controls, known as determinant risk factors for the morbidity-mortality of COPD.

In our study, legumes correlate with COPD status only in men but not in women or the whole population. Legumes are an essential component of a healthy diet and are associated with a reduced risk of COPD³³. This effect underlies "the Hispanic paradox"³³, relating to the anti-inflammatory effect of legumes (beans and lentils) on the reduction of systemic inflammation, decrease in respiratory symptoms, and eventually lower risk of COPD. Similarly, a low COPD risk has been reported in Asian subjects attributed to increased soy-product consumption³⁴. Legumes are found in most Moroccan dishes and are consumed as frequently as cereals³⁵. However, household legumes consumption has

been declining for years³⁵. This fact was confirmed through the current study, where the frequency of their consumption was less than once per week for low and moderate-MDA categories and twice a week for the high-MDA category (1.91 (\pm 0.95) for men and 1.86 (\pm 1.06) for women). The decreased consumption of legumes, mainly in women, may explain the absence of the association with COPD in this study.

Olive oil remains the primary source of dietary fat in Morocco and is incorporated in almost all meals³⁵. Our research has shown a decreased COPD risk with higher olive-oil consumption only in men, especially among the high-MDA group. This result may be due to the possible impact of phenolics which affects markers of oxidation and plasma susceptibility to oxidation caused by smoking. Although there is no direct causal relation between olive oil on pulmonary function, there is evidence of a particularly positive effect on improving respiratory functions, such as weight loss, lowering of triglycerides, lipoprotein cholesterol levels, and reduction of blood pressure³⁶. For women, and like other studies in the Mediterranean area, no association between olive oil consumption and COPD risk has been found³⁷, which can be explained by food component interactions during the digestion process.

There were also no statistically significant associations between fish intake and COPD risk. This result was in line with several prospective and longitudinal studies^{30,38}. The low consumption of fish may explain our results in the current sample (less than two times a week, which is lower than the recommendations of the MD pyramid²¹, and this is because the city of Fez is far away from the sea, and the consumption of fish which is not always affordable is not very widespread. Aside from the nutritional values of fish, its consumption is recognized as protective against chronic conditions due to omega 3 and 6, well known for their antioxidant and anti-inflammatory effects³⁹. These unsaturated fatty acids are crucial in restoring respiratory functions, reducing inflammation, and the burden of chronic diseases³⁰.

Little information about the relationship between dairy intake and lung function is known, especially among COPD patients⁴⁰. In the current study, a significantly positive association between the consumption of dairy products and COPD was found in the whole population and remained the same in the high-MDA category and women. This correlation can be explained by the high-fat content of dairy products eaten more frequently by COPD women (7.39 (\pm 4.73) in Men and 8.94 (\pm 6.16) in Women

$P < 0.001$). Other studies have shown similar results^{7,41}. In addition, data-driven dietary trends have shown the adverse impact of a diet rich in high-fat dairy products on respiratory symptoms⁷. Interestingly, it has been shown that the high-fat content of dairy products has been linked to worse respiratory symptoms and lower lung function. Conversely, higher intakes of low-fat dairy products may have a protective effect on lung function, likely through their anti-inflammatory action⁴⁰.

Our results on red meat and its derivatives on the deterioration of respiratory function were consistent with those of previous studies^{42,43}. In some studies, diets high in red meat and processed meat were associated with impaired respiratory functions (particularly FEV) and a high prevalence of COPD⁴⁴. This happens due to salts and nitrites in meats amplifying the nitrosative stress through the production of nitrogenous reagents, thus contributing to the deterioration of the progression of the pulmonary function. A Chinese cohort found that in COPD, and independently of a diet rich in fiber, diets rich in meat, sodium, and refined carbohydrates, could increase the risk of developing a cough with phlegm.

Although the results might be disaggregated by sex, the underlying data found that COPD risk was only related to men in moderate to high-MDA groups. The high smoking prevalence can explain the gender difference in the high COPD risk among men. Evidence has shown that smoking induces lung function impairment and poor health status⁴⁵. Moreover, smokers tend to adopt an unhealthy diet and consequently have a higher level of oxidative stress²³. These gender differences may also extend to different food choices and healthy lifestyles¹².

Our study had certain potential limitations, which must be considered. The first one is related to its cross-sectional nature in which the causal relations cannot be determined. The second limitation is linked to the nutritional data collection tool. Although the FFQ used in this study is valid and reliable, it collects data from the last 12 months preceding the interview. Thus, measurement errors could be introduced in the self-declared information. To improve the objectivity of the answers and reduce the possibility of misreporting, the FFQ was administered by trained interviewers who encouraged and helped participants to be more objective. In addition, since the FFQ represents a usual diet, nutritional estimates may be affected by seasonal changes and the risk of underestimating energy intake. However, this FFQ demonstrated correct validity and reproducibility for most nutrients. An advantage of these studies is that the spirometry test was performed according to the

BOLD protocol.

Furthermore, the quality of the spirometry tests and the database management system were checked and purified from non-compliant tests and data by the "Respiratory Epidemiology and Public Health Center at Imperial College." Several measurements were considered, including the forced expiratory volume (FEV₁), the most important measurement of lung function; it helps assess which level of COPD the patient has entered and gives more accuracy to COPD diagnosis and stages. Also, anthropometric measurements were collected using calibrated materials. Thus, the results could be considered accurate, and a wide range of potential confounding factors was controlled for in the analyses.

Conclusion

This study has followed a complementary approach to assess the association between COPD risk and MD components. It has confirmed previous results showing the significant association of the COPD risk with some components of MDS. However, since diet remains a modifiable lifestyle factor that may contribute to the prevention of chronic diseases like COPD, further extensive studies are needed to explore all MDS components better and shed light on more effective interventions in maintaining healthy eating habits and reducing COPD risk.

Conflict of interest

The authors declare that they have no competing interests.

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List of abbreviations

ATS: American Thoracic Society.

BMI: Body Mass Index.

BOLD: Burden of Obstructive Lung Disease

CI: Confident Interval.

COPD: Chronic Obstructive Pulmonary Disease.

ERS: European respiratory society.

FEV: Forced Expiratory Volume

FEV1: Forced Expiratory Volume in one second

FEV6: Forced Expiratory Volume in six second

FFQ: Food Frequency Questionnaire.

FVC: Forced Vital Capacity

GOLD: Global Initiative for Chronic Obstructive Lung Disease

MD: Mediterranean diet.

MDA: Mediterranean diet Adhearence.

MDS: Mediterranean diet Score.

MFCT: Moroccan food composition table.

OR: Odds-Ratio

SD: Standard Deviation.

TEI: Total Energy Intake

WC.: Waist Circumference.

WHO: World Health Organization

WHR: Waist to Hip Ratio

References

1. RGPH 2014 | Téléchargements | Site institutionnel du Haut-Commissariat au Plan du Royaume du Maroc. Accessed June 9, 2020. https://www.hcp.ma/downloads/RGPH-2014_t17441.html
2. El Rhazi K, Nejjari C, Romaguera D, et al. Adherence to a Mediterranean diet in Morocco and its correlates: cross-sectional analysis of a sample of the adult Moroccan population. *BMC Public Health*. 2012;12:345. doi:10.1186/1471-2458-12-345
3. Morocco country nutrition profile. Global Nutrition Report. Published 17:02:03.175006+00:00. Accessed May 27, 2019. <https://globalnutritionreport.org/nutrition-profiles/africa/northern-africa/morocco/>
4. Rabe KF, Hurd S, Anzueto A, et al. Global strategy for the diagnosis, management, and prevention of chronic obstructive pulmonary disease: GOLD executive summary. *Am J Respir Crit Care Med*. 2007;176(6):532-555. doi:10.1164/rccm.200703-456SO
5. El Rhazi K, Nejjari C, BenJelloun MC, El Biaze M, Attassi M, Garcia-Larsen V. Prevalence of chronic obstructive pulmonary disease in Fez, Morocco: results from the BOLD study. *Int J Tuberc Lung Dis Off J Int Union Tuberc Lung Dis*. 2016;20(1):136-141. doi:10.5588/ijtld.15.0029
6. Duffy S, Barnett S, Civic B, Criner GJ, A James Mamary MD, others. Risk of death by comorbidity prompting rehospitalization following the initial COPD hospitalization. *Chronic Obstr Pulm Dis J COPD Found*. 2015;2(1):17-22.
7. Scoditti E, Massaro M, Garbarino S, Toraldo DM. Role of Diet in Chronic Obstructive Pulmonary Disease Prevention and Treatment. *Nutrients*. 2019;11(6). doi:10.3390/nu11061357
8. Fischer A, Johansson I, Blomberg A, Sundström B. Adherence to a Mediterranean-like Diet as a Protective Factor Against COPD: A Nested Case-Control Study. *COPD*. 2019;16(3-4):272-277. doi:10.1080/15412555.2019.1634039
9. Trichopoulou A, Costacou T, Bamia C, Trichopoulos D. Adherence to a Mediterranean diet and survival in a Greek population. *N Engl J Med*. 2003;348(26):2599-2608. doi:10.1056/NEJMoa025039
10. Kaluza J, Harris HR, Linden A, Wolk A. Long-term consumption of fruits and vegetables and risk of chronic obstructive pulmonary disease: a prospective cohort study of women. *Int J Epidemiol*. 2018;47(6):1897-1909. doi:10.1093/ije/dyy178
11. Kaluza J, Harris H, Linden A, Wolk A. Long-term unprocessed and processed red meat consumption and risk of chronic obstructive pulmonary disease: a prospective cohort study of women. *Eur J Nutr*. 2019;58(2):665-672. doi:10.1007/s00394-018-1658-5
12. Gutiérrez-Carrasquilla L, Sánchez E, Hernández M, et al. Effects of Mediterranean Diet and Physical Activity on Pulmonary Function: A Cross-Sectional Analysis in the ILERVAS Project. *Nutrients*. 2019;11(2). doi:10.3390/nu11020329
13. El Rhazi K, Nejjari C, Zidouh A, Bakkali R, Berraho M, Barberger Gateau P. Prevalence of obesity and associated sociodemographic and lifestyle factors in Morocco. *Public Health Nutr*. 2011;14(1):160-167. doi:10.1017/S1368980010001825
14. Buist AS, Vollmer WM, Sullivan SD, et al. The Burden of Obstructive Lung Disease Initiative (BOLD): rationale and design. *COPD*. 2005;2(2):277-283.
15. World Health Organization. *Waist Circumference and Waist-Hip Ratio: Report of a WHO Expert Consultation, Geneva, 8-11 December 2008*. World Health Organization; 2011.
16. Enright P, Vollmer WM, Lamprecht B, et al. Quality of spirometry tests performed by 9893 adults in 14 countries: the BOLD Study. *Respir Med*. 2011;105(10):1507-1515.
17. Vogelmeier CF, Criner GJ, Martinez FJ, et al. Global Strategy for the Diagnosis, Management, and Prevention of Chronic Obstructive Lung Disease 2017 Report. GOLD Executive Summary. *Am J Respir Crit Care Med*. 2017;195(5):557-582. doi:10.1164/rccm.201701-0218PP
18. El Kinany K, Garcia-Larsen V, Khalis M, et al. Adaptation and validation of a food frequency questionnaire (FFQ) to assess dietary intake in Moroccan adults. *Nutr J*. 2018;17(1):61. doi:10.1186/s12937-018-0368-4

19. Khalis M, Garcia-Larsen V, Charaka H, et al. Update of the Moroccan food composition tables: Towards a more reliable tool for nutrition research. *J Food Compos Anal.* 2020;87:103397. doi:10.1016/j.jfca.2019.103397
20. Thiébaud A, Kesse E, Com-Nougé C, Clavel-Chapelon F, Bénichou J. Ajustement sur l'apport énergétique dans l'évaluation des facteurs de risque alimentaires. *Rev D'Épidémiologie Santé Publique.* 2004;52(6):539-557. doi:10.1016/S0398-7620(04)99093-1
21. Bach-Faig A, Berry EM, Lairon D, et al. Mediterranean diet pyramid today. Science and cultural updates. *Public Health Nutr.* 2011;14(12A):2274-2284. doi:10.1017/S1368980011002515
22. Serra-Majem L, Tomaino L, Dernini S, et al. Updating the Mediterranean Diet Pyramid towards Sustainability: Focus on Environmental Concerns. *Int J Environ Res Public Health.* 2020;17(23):8758. doi:10.3390/ijerph17238758
23. Martín-Luján F, Catalin RE, Salamanca-González P, et al. A clinical trial to evaluate the effect of the Mediterranean diet on smokers lung function. *NPJ Prim Care Respir Med.* 2019;29(1):40. doi:10.1038/s41533-019-0153-7
24. Papanikolaou Y, Fulgoni VL. Certain Grain Foods Can Be Meaningful Contributors to Nutrient Density in the Diets of U.S. Children and Adolescents: Data from the National Health and Nutrition Examination Survey, 2009-2012. *Nutrients.* 2017;9(2). doi:10.3390/nu9020160
25. Veronese N, Solmi M, Caruso MG, et al. Dietary fiber and health outcomes: an umbrella review of systematic reviews and meta-analyses. *Am J Clin Nutr.* 2018;107(3):436-444. doi:10.1093/ajcn/nqx082
26. Kan H, Stevens J, Heiss G, Rose KM, London SJ. Dietary fiber, lung function, and chronic obstructive pulmonary disease in the atherosclerosis risk in communities study. *Am J Epidemiol.* 2008;167(5):570-578. doi:10.1093/aje/kwm343
27. Varraso R, Willett WC, Camargo CA. Prospective Study of Dietary Fiber and Risk of Chronic Obstructive Pulmonary Disease Among US Women and Men. *Am J Epidemiol.* 2010;171(7):776-784. doi:10.1093/aje/kwp455
28. Tabak C, Smit HA, Heederik D, Ocké MC, Kromhout D. Diet and chronic obstructive pulmonary disease: independent beneficial effects of fruits, whole grains, and alcohol (the MORGEN study). *Clin Exp Allergy J Br Soc Allergy Clin Immunol.* 2001;31(5):747-755. doi:10.1046/j.1365-2222.2001.01064.x
29. Zhai H, Wang Y, Jiang W. Fruit and Vegetable Intake and the Risk of Chronic Obstructive Pulmonary Disease: A Dose-Response Meta-Analysis of Observational Studies. *BioMed Res Int.* 2020;2020:3783481. doi:10.1155/2020/3783481
30. Varraso R, Barr RG, Willett WC, Speizer FE, Camargo CA. Fish intake and risk of chronic obstructive pulmonary disease in 2 large US cohorts. *Am J Clin Nutr.* 2015;101(2):354-361. doi:10.3945/ajcn.114.094516
31. Garcia-Larsen V, Potts JF, Omenaas E, et al. Dietary antioxidants and 10-year lung function decline in adults from the ECRHS survey. *Eur Respir J.* 2017;50(6). doi:10.1183/13993003.02286-2016
32. Kaluza J, Larsson SC, Orsini N, Linden A, Wolk A. Fruit and vegetable consumption and risk of COPD: a prospective cohort study of men. *Thorax.* 2017;72(6):500-509. doi:10.1136/thoraxjnl-2015-207851
33. Young RP, Hopkins RJ. A review of the Hispanic paradox: time to spill the beans? *Eur Respir Rev Off J Eur Respir Soc.* 2014;23(134):439-449. doi:10.1183/09059180.00000814
34. Hirayama F, Lee AH, Binns CW, et al. Soy consumption and risk of COPD and respiratory symptoms: a case-control study in Japan. *Respir Res.* 2009;10:56. doi:10.1186/1465-9921-10-56
35. Enquête Nationale sur la Consommation et les Dépenses des Ménages | Téléchargements | Site institutionnel du Haut-Commissariat au Plan du Royaume du Maroc. Accessed May 3, 2020. https://www.hcp.ma/downloads/Enquete-Nationale-sur-la-Consommation-et-les-Depenses-des-Menages_t21181.html
36. Ray NB, Hilsabeck KD, Karagiannis TC, McCord DE. Chapter 36 - Bioactive Olive Oil Polyphenols in the Promotion of Health. In: Singh RB, Watson RR, Takahashi T, eds. *The Role of Functional Food Security in Global*

- Health. Academic Press; 2019:623-637. doi:10.1016/B978-0-12-813148-0.00036-0*
37. Mattioli V, Zanolin ME, Cazzoletti L, et al. Dietary flavonoids and respiratory diseases: a population-based multi-case–control study in Italian adults. *Public Health Nutr.* 2020;23(14):2548-2556. doi:10.1017/S1368980019003562
 38. Walda IC, Tabak C, Smit HA, et al. Diet and 20-year chronic obstructive pulmonary disease mortality in middle-aged men from three European countries. *Eur J Clin Nutr.* 2002;56(7):638-643. doi:10.1038/sj.ejcn.1601370
 39. de Batlle J, Sauleda J, Balcells E, et al. Association between $\Omega 3$ and $\Omega 6$ fatty acid intakes and serum inflammatory markers in COPD. *J Nutr Biochem.* 2012;23(7):817-821. doi:10.1016/j.jnutbio.2011.04.005
 40. Jiang R, Jacobs DR, He K, et al. Associations of dairy intake with CT lung density and lung function. *J Am Coll Nutr.* 2010;29(5):494-502.
 41. Butler LM, Koh WP, Lee HP, Tseng M, Yu MC, London SJ. Prospective Study of Dietary Patterns and Persistent Cough with Phlegm among Chinese Singaporeans. *Am J Respir Crit Care Med.* 2006;173(3):264-270. doi:10.1164/rccm.200506-901OC
 42. Salari-Moghaddam A, Milajerdi A, Larijani B, Esmailzadeh A. Processed red meat intake and risk of COPD: A systematic review and dose-response meta-analysis of prospective cohort studies. *Clin Nutr Edinb Scotl.* 2019;38(3):1109-1116. doi:10.1016/j.clnu.2018.05.020
 43. de Batlle J, Mendez M, Romieu I, et al. Cured meat consumption increases risk of readmission in COPD patients. *Eur Respir J.* 2012;40(3):555-560. doi:10.1183/09031936.00116911
 44. Jiang R, Paik DC, Hankinson JL, Barr RG. Cured meat consumption, lung function, and chronic obstructive pulmonary disease among United States adults. *Am J Respir Crit Care Med.* 2007;175(8):798-804. doi:10.1164/rccm.200607-969OC
 45. Bartal M. COPD and tobacco smoke. *Monaldi Arch Chest Dis Arch Monaldi Mal Torace.* 2005;63(4):213-225. doi:10.4081/monaldi.2005.623