



Association between dietary antioxidant intakes and chronic respiratory diseases in adults

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ABSTRACT

Background: Chronic respiratory diseases (CRDs) pose a significant global health burden. Antioxidant-rich diets have been associated with improved lung health, but the specific relationship with CRDs remains unclear.

Methods: This study examined the relationship between dietary antioxidant intakes and CRDs using data from the 2001–2018 National Health and Nutrition Examination Survey (NHANES). Information on dietary antioxidant intakes, including vitamins A, C, and E, zinc, selenium, and carotenoid, were collected from the 2 24-h recall interviews to calculate composite dietary antioxidant index (CDAI). CRDs were determined based on self-reported physician diagnoses. To examine the relationship between CDAI and CRDs, multivariate logistic regression was used. To study potential non-linear correlations within these associations, restricted cubic spline (RCS) regression was performed.

Results: The study involved 40 557 individuals. The median CDAI was -0.09 ($-2.05, 2.25$). We discovered those who were in the fourth quartile of CDAI scores had a 19% lower prevalence than those in the first quartile ($OR = 0.81$ [$0.72-0.91$], $P_{trend} < 0.01$) after adjusting for all relevant covariates. The fourth quartile of CDAI was linked with a lower prevalence of emphysema ($OR = 0.57$ [$0.40-0.81$], $P_{trend} < 0.01$) and chronic bronchitis ($OR = 0.74$ [$0.62-0.88$], $P_{trend} < 0.01$). RCS regression showed that CDAI was non-linearly related to the prevalence of CRDs, with inflection points of 3.20 (P for non-linearity < 0.01). The stratified analysis did not identify variables that significantly affected the results.

Conclusion: Higher dietary antioxidant intakes were related with a lower prevalence of CRDs (particularly emphysema and chronic bronchitis) in general adults.

Keywords: CRDs, Antioxidants, Dietary intakes, NHANES

INTRODUCTION

Chronic respiratory diseases (CRDs) encompass various conditions such as asthma, emphysema,

and chronic bronchitis, which significantly impair lung function and overall respiratory health.¹ Asthma is characterized by recurrent episodes of

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coughing, chest tightness, and shortness of breath.² Emphysema is a progressive lung disease that damages and destroys the air sacs in the lungs.³ Chronic bronchitis involves airway inflammation and constriction, leading to excessive mucus production and persistent cough.⁴ These diseases pose a substantial global burden of morbidity and present a significant challenge to the global healthcare system. The development of CRDs is influenced by various factors, including environmental exposures such as air pollution, smoking, occupational hazards, and genetic predispositions.⁵⁻⁷ Additionally, lifestyle factors like physical inactivity and poor nutrition contribute to the severity of CRDs.⁸⁻¹⁰ Epidemiological research suggests that dietary variables, such as antioxidants, may play a vital role in the prevention and management of CRDs.^{11,12}

The Composite Dietary Antioxidant Index (CDAI) is a metric used to assess the total antioxidant capacity of an individual's diet.¹³ It combines multiple dietary antioxidants, such as vitamins (A, C, E), minerals (selenium, zinc), and phytochemicals (carotenoids), into a single index. By evaluating the collective impact of multiple antioxidants, the CDAI provides a comprehensive measure of the diet's ability to combat oxidative stress.¹⁴ Imbalance between free radicals and antioxidants in the body leads to cellular damage.¹⁵ Studies have demonstrated that antioxidant-rich diets can reduce the risk of developing chronic diseases, including depression, osteoporosis, cancer, and cardiovascular mortality.¹⁶⁻¹⁹ Antioxidants play a protective role in human health through various mechanisms.²⁰ They can directly scavenge reactive oxygen species (ROS) and prevent them from causing damage to body tissues.²¹ Additionally, they upregulate endogenous antioxidant enzymes to enhance the body's defenses against oxidative stress.²² By considering both the quantity of antioxidants consumed, the CDAI offers a comprehensive perspective on their potential health benefits.

Several studies have suggested that dietary antioxidants may benefit respiratory health. One meta-analysis reported that increased dietary antioxidant intake was associated with a reduced likelihood of developing chronic obstructive pulmonary disease (COPD).²³ Another study

focusing on asthmatics found that increasing dietary antioxidant intake may help reduce the frequency of asthma symptoms.²⁴ However, few studies have specifically investigated the relationship between the CDAI and the onset or progression of CRDs. Thus, a comprehensive investigation of this association is warranted. Our study aims to explore the potential protective role of dietary antioxidants against CRDs in adults by analyzing data collected from a representative population. We examined CDAs obtained from the 2001-2018 National Health and Nutrition Examination Survey (NHANES) to investigate the link between CDAs and the prevalence of CRDs. By exploring the correlation between dietary antioxidant intake and respiratory health, we aim to identify potential modifiable factors that could contribute to the development of tailored dietary recommendations for individuals susceptible to CRDs.

MATERIALS AND METHODS

Study population

The NHANES is a program conducted by the Centers for Disease Control and Prevention (CDC) in the United States.²⁵ It collects comprehensive health and nutritional data from a nationally representative population. The survey collects information through interviews, medical exams, and laboratory tests, providing valuable data on various aspects of health. The research protocols were approved by the National Center for Health Statistics (NCHS) Research Ethics Review Board, and all participants provided informed permission.

In total, 91 351 people took part in the NHANES from 2001 to 2018. We excluded 41 150 participants with an age <20 and missing information on CRDs. A total of 5700 participants without data to calculate CDAI, 1148 participants who were pregnant and 2796 participants with extreme energy intakes were also ruled out. We eventually included 40 557 participants for analyses (Fig. 1).

Assessment of CDAI and CRDs

In the NHANES, dietary intake data were acquired through a structured 24-h dietary recall interview. The NHANES employs a computer-assisted dietary interview system,

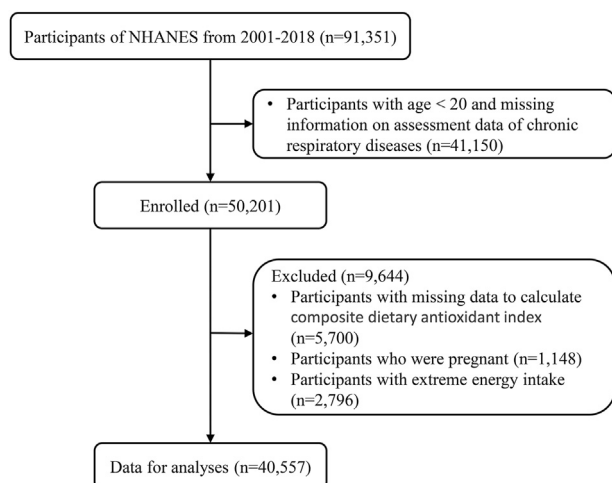


Fig. 1 Flowchart of the study participants.

which systematically gathers and records details concerning the variety and quantities of ingestible items, encompassing food, beverages (inclusive of water), consumed during the 24 h preceding the interviews. The initial dietary recall interview was administered in a face-to-face setting at the Mobile Examination Center (MEC), while the subsequent interview was conducted via telephone within a period ranging from 3 to 10 days after the first encounter. The assessment of dietary nutrient intake and total energy consumption relied on the University of Texas Food Intake Analysis System in conjunction with data sourced from the United States Department of Agriculture Survey Nutrient Database. To obtain information regarding the intake of six specific dietary antioxidants as well as total energy intake, a process of averaging the results from two separate 24-h recall interviews was undertaken. It is noteworthy that the nutritional assessments did not encompass any nutrients originating from dietary supplements or medicinal compounds.

The CDAI is a calculating method for determining the whole antioxidant amount found in a person's diet.¹³ It contains 6 dietary antioxidants – vitamins A, C, and E, as well as zinc, selenium, and carotenoids – and assigns weights to each antioxidant based on its potential health benefits.²⁶ To estimate the CDAI, we standardized each of the same six dietary vitamins and minerals by subtracting the global average and dividing by the global standard deviation. We then calculated the CDAI by

adding the standard intake of these vitamins. Higher CDAI scores indicate diets that contain more antioxidants and types of antioxidants. In this study, the effect of abnormal values was avoided by excluding participants with abnormal total energy intake (>4200 or <800 kcal/day in male; >3500 or <500 kcal/day in female).

CRDs refer to long-term conditions affecting the airways and lungs, including asthma, emphysema, and chronic bronchitis.⁵ People with CRDs were defined as affirming the following question: Has a doctor or other health professional ever told you that you have asthma/emphysema/chronic bronchitis?

Covariates

Information on age (years), sex (male or female), race/ethnicity (Mexican American, other Hispanic, non-Hispanic White, non-Hispanic Black, or other), education level (below high school, high school, or above high school), total energy intake (kcal/day), and carotenoid supplement use (%) was collected through a standardized approach. The poverty income ratio (PIR) is derived by dividing the household's earnings by a factor unique to the size and composition of the household and categorised into 3 groups (≤ 1.0 , 1.1–3.0, or > 3.0).²⁷ Individual smoking and drinking status were recorded through a standardised questionnaire asking participants about their past and present smoking and drinking status.²⁸ Smoking status was classified as never smokers (<100 cigarettes), current smokers (>100 cigarettes), and former smokers (>100 cigarettes and had quit smoking). Drinking status was classified as nondrinker, low-to-moderate drinker (<1 drink/day for females and <2 drinks/day for males), or heavy drinker (≥ 1 drink/day for females and ≥ 2 drinks/day for males). Physical activity was classified as inactive (no leisure-time physical activity), insufficiently active (moderate activity 1–5 times per week with metabolic equivalents [MET] 3–6 or vigorous activity 1–3 times per week with MET > 6), and active (moderate activity > 5 times per week with MET 3–6 or vigorous activity > 3 times per week with MET > 6).^{29,30} Self-reported questionnaires were used to collect information on the prevalence of hypertension and diabetes.

Statistical analysis

The individuals' initial features are expressed as means with standard errors (SEs), medians with interquartile ranges (IQRs), or numbers with percentages. The Student's *t*-test or Mann-Whitney *U* test were used to compare continuous variables, while the chi-square test was used to compare categorical variables. The concentration and distribution of CDAI and its components were analysed. The association coefficients between serum and dietary antioxidant micronutrients were determined using pairwise Spearman correlation analysis. The study population was categorised according to CDAI quartiles.

Multifactorial logistic regression was used to analyse the link between CDAI and its components with the prevalence of CRDs, asthma, emphysema and chronic bronchitis in the United States (US) population. Potential non-linear links between CDAI and prevalence of CRDs were analysed using restricted cubic spline (RCS) regression model with 10th, 50th and 90th percentile as nodes. RCS is a flexible and non-linear regression approach that allows for a curved relationship by fitting piecewise polynomials. Stratified analyses were used to explore the effect of different characteristics on the associations of CDAI and its components with the prevalence of CRDs, including age (20–39, 40–59, or ≥ 60 years), sex (female, or male), race (non-Hispanic White, or other), family PIR (≤ 1.0 , 1.1–3.0, or > 3.0), smoking status (never, former, or current smoker), drinking status (nondrinker, low-to-moderate drinker, or heavy drinker), physical activity (inactive, insufficiently active, or active), BMI (< 25.0 , 25.0–29.9, or > 29.9 kg/m²), supplement use (no, or yes). All analyses were performed using R (version 4.2.0) and *P*-values less than 0.05 were considered statistically significant.

RESULTS

Baseline characteristics of the participants

Table 1 shows the NHANES initial features for the general adults from 2001 to 2018. A total of 40 557 adults (7110 participants with CRDs) were included in this analysis. The participants were 48.42% male and 45.23% non-Hispanic white. Except for education level and drinking status ($P > 0.05$), all baseline variables were significantly

different between people with and without CRDs. Patients with CRDs were more likely to be old women (age ≥ 60 years), female, non-Hispanic white, current smokers, obese, and physically inactive ($P < 0.01$). They were also more probable to have hypertension and diabetes ($P = 0.01$). We also looked at the research population's baseline characteristics based on CDAI quartiles (Table S2). We found higher proportions of middle-aged women (age 40–59 years), non-smokers, low-to-moderate drinkers, normal-weight individuals, and physically active individuals among participants in the fourth quartile of the CDAI ($P < 0.01$). They tended to be less prone to self-reported hypertension and diabetes ($P < 0.01$).

Distributions and concentrations of CDAI among adults with CRDs

Table S1 listed the distribution and concentration of CDAI and its components among adults with CRDs. The mean CDAI was -0.09 (-2.05 , 2.25). The average intake of vitamins A, C, and E, and zinc, selenium, and carotenoids in the study population was approximately 481.00 (270.00, 785.00) $\mu\text{g/day}$, 51.50 (21.50, 111.60) mg/day, 6.58 (4.29, 9.91) mg/day, 9.93 (6.89, 14.08) mg/day, 99.60 (70.10, 136.80) $\mu\text{g/day}$, and 5344.00 (2002.00, 12162.00) $\mu\text{g/day}$, respectively. Figure S1. showed the Spearman correlation coefficients among serum and dietary antioxidant micronutrients. Significant correlations were found between both CDAI and individual dietary antioxidant micronutrients ($r > 0.55$). A strong correlation could be observed between serum zinc and serum arsenic ($r = 0.63$).

Association between CDAI and CRDs

We found a substantial negative connection between CDAI scores and the frequency of CRDs in all models (Table 2). Higher CDAI scores were linked with a lower prevalence of CRDs. After controlling for all variables, individuals in the fourth quartile of CDAI scores had a 19% lower prevalence than those in the first quartile (OR = 0.81 [0.72–0.91], $P_{\text{trend}} < 0.01$). In the unadjusted model and model 2, CDAI was found to be negatively associated with the prevalence of emphysema and chronic bronchitis. Compared to the first quartile, the fourth quartile of CDAI was negatively associated

Characteristics	Total (n = 40 557)	Chronic respiratory diseases		P value
		No (n = 33 447)	Yes (n = 7110)	
Age, years				<0.01
20-39	12 884 (31.77)	10 654 (35.44)	2230 (35.43)	
40-59	13 281 (32.75)	11 051 (38.81)	2230 (35.87)	
≥60	14 392 (35.49)	11 742 (25.75)	2650 (28.70)	
Sex, %				<0.01
Female	20 921 (51.58)	16 807 (50.93)	4114 (59.57)	
Male	19 636 (48.42)	16 640 (49.07)	2996 (40.43)	
Race/ethnicity, %				<0.01
Mexican American	6658 (16.42)	6001 (8.76)	657 (4.46)	
Other Hispanic	3385 (8.35)	2788 (5.34)	597 (4.97)	
Non-Hispanic White	18 345 (45.23)	14 606 (68.29)	3739 (73.19)	
Non-Hispanic Black	8439 (20.81)	6862 (10.65)	1577 (11.25)	
Other race	3730 (9.2)	3190 (6.96)	540 (6.12)	
Education level, %				0.74
Below high school	10 311 (25.42)	8594 (15.99)	1717 (16.43)	
High school	9425 (23.24)	7750 (23.83)	1675 (23.98)	
Above high school	20 821 (51.34)	17 103 (60.17)	3718 (59.59)	
Family PIR, %				<0.01
≤1.0	8219 (20.27)	6521 (13.01)	1698 (17.18)	
1.1-3.0	17 097 (42.16)	14 032 (35.46)	3065 (38.11)	
>3.0	15 241 (37.58)	12 894 (51.52)	2347 (44.71)	
Smoking status, %				<0.01
Never smoker	22 066 (54.41)	18 861 (56.31)	3205 (45.73)	
Former smoker	10 216 (25.19)	8176 (24.43)	2040 (28.13)	
Current smoker	8275 (20.4)	6410 (19.26)	1865 (26.14)	
Drinking status, %				0.19
Nondrinker	9370 (23.1)	7789 (18.82)	1581 (18.74)	
Low-to-moderate drinker	28 020 (69.09)	23 038 (71.86)	4982 (72.80)	
Heavy drinker	3167 (7.81)	2620 (9.32)	547 (8.46)	
Body mass index, %				<0.01
<25.0 kg/m ²	11 674 (28.78)	9864 (30.91)	1810 (27.42)	
25.0-29.9 kg/m ²	13 680 (33.73)	11 622 (34.26)	2058 (28.40)	
>29.9 kg/m ²	15 203 (37.49)	11 961 (34.84)	3242 (44.18)	
Physical activity, %				<0.01
Inactive	10 991 (27.1)	8877 (21.22)	2114 (24.93)	
Insufficiently active	14 885 (36.7)	12 396 (39.79)	2489 (37.74)	
Active	14 681 (36.2)	12 174 (38.98)	2507 (37.34)	
Total energy intakes, kcal/day				0.01
Quartile 1	10 148 (25.02)	8212 (21.42)	1936 (23.65)	
Quartile 2	10 136 (24.99)	8351 (24.29)	1785 (24.34)	
Quartile 3	10 136 (24.99)	8472 (26.14)	1664 (24.76)	
Quartile 4	10 137 (24.99)	8412 (28.15)	1725 (27.26)	
Self-reported hypertension, %				<0.01
No	22 955 (56.6)	19 422 (63.57)	3533 (55.66)	
Yes	17 602 (43.4)	14 025 (36.43)	3577 (44.34)	

(continued)

Characteristics	Total (n = 40 557)	Chronic respiratory diseases		P value
		No (n = 33 447)	Yes (n = 7110)	
Self-reported diabetes, %				<0.01
No	35 371 (87.21)	29 450 (91.33)	5921 (87.47)	
Yes	5186 (12.79)	3997 (8.67)	1189 (12.53)	
Supplement use, %				0.01
No	19 920 (49.12)	16 618 (46.31)	3302 (44.19)	
Yes	20 637 (50.88)	16 829 (53.69)	3808 (55.81)	

Table 1. (Continued) Baseline characteristics of the general adult population in NHANES 2001–2018. Abbreviations: PIR, poverty income ratio. Categorical variables are presented as numbers (percentages). Sampling weights were applied for calculation of demographic descriptive statistics; N reflect the study sample while percentages reflect the survey-weighted data

with the prevalence of emphysema (OR = 0.57 [0.40–0.81], $P_{\text{trend}} < 0.01$) and chronic bronchitis (OR = 0.74 [0.62–0.88], $P_{\text{trend}} < 0.01$) after adjusting for all confounders. However, no association was observed between CDAI scores and the prevalence of asthma. Furthermore, we investigated the relationship between each dietary antioxidant micronutrient quartile and the occurrence of CRDs in adults (Table S3).

The dose-response relationship between CDAI and CRDs was further analysed using RCS (Fig. 2). We found a non-linear negative correlation between CDAI and the prevalence of CRDs (P for nonlinearity < 0.01), with an inflection point of 3.20. This non-linear relationship was also present between CDAI and the prevalence of chronic bronchitis (P for nonlinearity < 0.01), with an inflection point of 4.33. However, there was a negative linear

	Quartiles of CDAI				P_{trend}
	<-2.26	[-2.26 to -0.37]	[-0.38-1.98]	>1.98	
CRDs					
Crude	1 [Reference]	0.87 (0.79-0.96)	0.80 (0.72-0.88)	0.81 (0.73-0.89)	<0.01
Model 1	1 [Reference]	0.90 (0.82-0.99)	0.82 (0.74-0.90)	0.83 (0.75-0.91)	<0.01
Model 2	1 [Reference]	0.91 (0.82-1.01)	0.82 (0.74-0.91)	0.81 (0.72-0.91)	<0.01
Asthma					
Crude	1 [Reference]	0.90 (0.82-1.00)	0.87 (0.79-0.96)	0.90 (0.80-1.02)	0.09
Model 1	1 [Reference]	0.94 (0.85-1.04)	0.90 (0.82-1.00)	0.93 (0.82-1.05)	0.19
Model 2	1 [Reference]	0.92 (0.82-1.02)	0.87 (0.79-0.97)	0.87 (0.76-1.00)	0.05
Emphysema					
Crude	1 [Reference]	0.86 (0.68-1.09)	0.59 (0.46-0.75)	0.44 (0.33-0.58)	<0.01
Model 1	1 [Reference]	0.81 (0.63-1.02)	0.56 (0.44-0.71)	0.43 (0.32-0.57)	<0.01
Model 2	1 [Reference]	0.96 (0.74-1.25)	0.72 (0.53-0.97)	0.57 (0.40-0.81)	<0.01
Chronic bronchitis					
Crude	1 [Reference]	0.77 (0.67-0.90)	0.63 (0.54-0.73)	0.63 (0.55-0.72)	<0.01
Model 1	1 [Reference]	0.80 (0.68-0.93)	0.64 (0.55-0.75)	0.64 (0.55-0.74)	<0.01
Model 2	1 [Reference]	0.86 (0.73-1.02)	0.73 (0.62-0.86)	0.74 (0.62-0.88)	<0.01

Table 2. ORs (95% CIs) of the prevalence of chronic respiratory diseases (CRDs) and its components according to quartiles of composite dietary antioxidant index (CDAI) among adults in NHANES 2001–2018. Model 1 was adjusted for age (20–39, 40–59, or ≥ 60), sex (male or female), and race (Mexican American, Other Hispanic, Non-Hispanic White, Non-Hispanic Black or Other). Model 2 was adjusted for Model 1 plus education level (below high school, high school, or above high school), family income-to-poverty ratio (≤ 1.0 , 1.1–3.0, or > 3.0), smoking status (never smoker, former smoker, or current smoker), drinking status (nondrinker, low-to-moderate drinker, or heavy drinker), BMI (< 25.0 , 25.0–29.9, or > 29.9), energy intake levels (in quartiles), physical activity (inactive, insufficiently active, or active), diabetes (yes or no), hypertension (yes or no), and supplement use (yes or no)

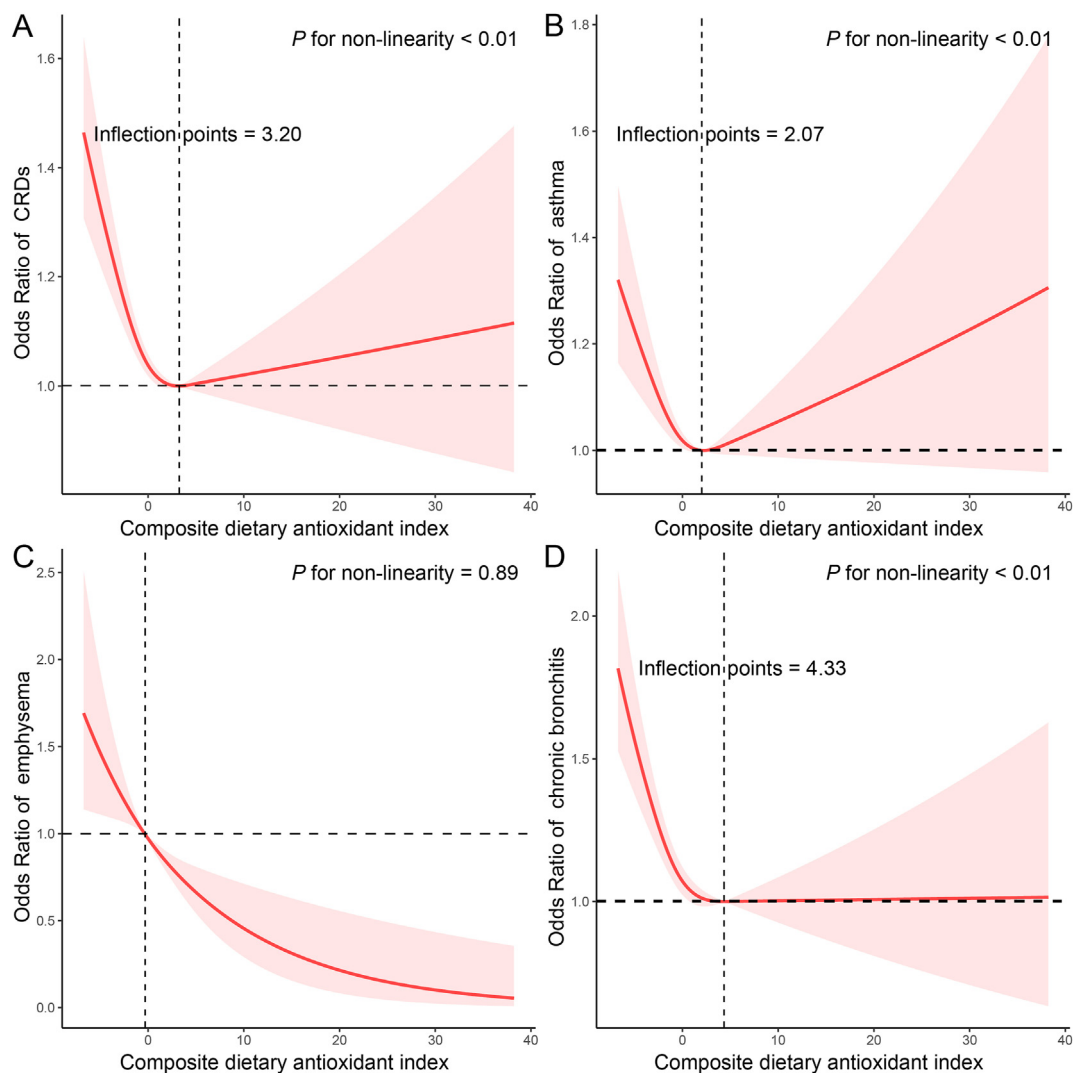


Fig. 2 The exposure-response association of the composite dietary antioxidant index (CDAI) with the prevalence of chronic respiratory diseases (CRDs) by restricted cubic spline (RCS). Analyses were adjusted for covariates age (20-39, 40-59, or ≥ 60), sex (male or female), race (Mexican American, Other Hispanic, Non-Hispanic White, Non-Hispanic Black or Other), education level (below high school, high school, or above high school), family income-to-poverty ratio (≤ 1.0 , 1.1-3.0, or > 3.0), smoking status (never smoker, former smoker, or current smoker), drinking status (nondrinker, low-to-moderate drinker, or heavy drinker), BMI (< 25.0 , 25.0-29.9, or > 29.9), energy intake levels (in quartiles), physical activity (inactive, insufficiently active, or active), diabetes (yes or no), hypertension (yes or no), and supplement use (yes or no).

association between CDAI and the prevalence of emphysema (P for nonlinearity = 0.89).

Stratified analysis

We analysed the study population in subgroups according to age, sex, race, family PIR, smoking and drinking status, physical activity, BMI, and supplement use (Table 3). Interestingly, we found that the association between CDAI and prevalence of CRDs was consistent across all subgroups, suggesting that it was not influenced by these demographic factors.

DISCUSSION

We conducted a study with a total of 40 557 participants to investigate the potential connection between CDAI scores and the prevalence of CRDs. Our findings revealed that individuals in the fourth quartile of CDAI scores exhibited a 19% lower prevalence of CRDs compared to those in the first quartile (OR = 0.81 [0.72-0.91], $P_{\text{trend}} < 0.01$). Among the 3 CRDs examined, CDAI showed a negative association with the prevalence of chronic bronchitis and emphysema, but no significant association was observed with asthma. Notably, this

Subgroups	N	Quartiles of CDAI				P _{interaction}
		<-2.26	[-2.26, -0.37]	[-0.38,1.98]	>1.98	
Age, years						
20-39	12 884	1 [Reference]	0.86 (0.73,1.01)	0.77 (0.65,0.92)	0.83 (0.69,0.99)	0.29
40-59	13 281	1 [Reference]	0.86 (0.72,1.03)	0.79 (0.65,0.95)	0.67 (0.55,0.83)	
≥60	14 392	1 [Reference]	1.00 (0.85,1.18)	0.90 (0.73,1.10)	0.97 (0.79,1.20)	
Sex, %						
Female	20 921	1 [Reference]	0.99 (0.84,1.17)	0.87 (0.73,1.05)	0.91 (0.75,1.10)	0.40
Male	19 636	1 [Reference]	0.86 (0.75,0.97)	0.79 (0.69,0.89)	0.75 (0.64,0.86)	
Race, %						
Non-Hispanic White	18 345	1 [Reference]	0.93 (0.81,1.07)	0.84 (0.73,0.97)	0.82 (0.70,0.96)	0.77
Other	22 212	1 [Reference]	0.83 (0.73,0.96)	0.77 (0.65,0.90)	0.78 (0.66,0.92)	
Family PIR, %						
≤1.0	8219	1 [Reference]	0.91 (0.74,1.11)	0.85 (0.69,1.05)	0.74 (0.59,0.93)	0.09
1.1-3.0	17 097	1 [Reference]	0.89 (0.77,1.04)	0.72 (0.62,0.84)	0.77 (0.66,0.89)	
>3.0	15 241	1 [Reference]	0.94 (0.79,1.11)	0.92 (0.78,1.09)	0.89 (0.73,1.10)	
Smoking status, %						
Never smoker	22 066	1 [Reference]	0.90 (0.78,1.05)	0.84 (0.72,0.97)	0.90 (0.76,1.06)	0.32
Former smoker	10 216	1 [Reference]	0.92 (0.76,1.10)	0.81 (0.65,1.02)	0.74 (0.59,0.92)	
Current smoker	8275	1 [Reference]	0.92 (0.76,1.13)	0.84 (0.69,1.03)	0.74 (0.59,0.92)	
Drinking status, %						
Nondrinker	9370	1 [Reference]	1.03 (0.83,1.29)	0.99 (0.77,1.26)	0.88 (0.69,1.12)	0.43
Low-to-moderate drinker	28 020	1 [Reference]	0.87 (0.76,0.98)	0.75 (0.66,0.86)	0.76 (0.65,0.88)	
Heavy drinker	3167	1 [Reference]	0.92 (0.65,1.29)	1.03 (0.71,1.50)	1.06 (0.70,1.61)	
Physical activity, %						
Inactive	10 991	1 [Reference]	0.98 (0.82,1.17)	0.80 (0.65,0.99)	0.96 (0.78,1.19)	0.65
Insufficiently active	14 885	1 [Reference]	0.87 (0.75,1.02)	0.79 (0.68,0.92)	0.73 (0.62,0.86)	
Active	14 681	1 [Reference]	0.90 (0.73,1.10)	0.85 (0.70,1.03)	0.81 (0.65,1.01)	
Body mass index, %						
<25.0 kg/m ²	11 674	1 [Reference]	0.96 (0.77,1.20)	0.90 (0.75,1.08)	0.82 (0.66,1.01)	0.80
25.0-29.9 kg/m ²	13 680	1 [Reference]	0.95 (0.79,1.14)	0.82 (0.67,1.00)	0.81 (0.66,0.99)	
>29.9 kg/m ²	15 203	1 [Reference]	0.86 (0.74,0.99)	0.78 (0.67,0.92)	0.82 (0.69,0.97)	

Supplement use, %							
No	19 920	1 [Reference]	0.88 (0.76,1.01)	0.80 (0.70,0.92)	0.77 (0.64,0.93)	0.99	
Yes	20 637	1 [Reference]	0.93 (0.82,1.06)	0.84 (0.72,0.98)	0.85 (0.72,0.99)		

Table 3. Stratified analyses of the associations between quartiles of composite dietary antioxidant index (CDAI) and the prevalence of chronic respiratory diseases (CRDs) in NHANES 2001-2018. Analyses were adjusted for covariates age (20-39, 40-59, or ≥60), sex (male or female), race (Mexican American, Other Hispanic, Non-Hispanic Black or Other), education level (below high school, high school, or above high school), family income-to-poverty ratio (≤1.0, 1.1-3.0, or >3.0), smoking status (never smoker, former smoker, or current smoker), drinking status (nondrinker, low-to-moderate drinker, or heavy drinker), BMI (<25.0, 25.0-29.9, or >29.9), energy intake levels (in quartiles), physical activity (inactive, insufficiently active, or active), diabetes (yes or no), hypertension (yes or no), and supplement use (yes or no) when they were not the strata variables

association between CDAI and CRDs remained consistent across all subgroups analysed. This pioneering study is the first to demonstrate a link between CDAI and CRD prevalence in the general population.

The CDAI is an indicator used to quantify total dietary antioxidant intake,¹³ taking into account antioxidants from various dietary sources such as fruits, vegetables, tea, and coffee.³¹ Previous studies have indicated that higher CDAI scores (indicating greater antioxidant intake) are associated with a reduced risk of developing diabetic nephropathy.³² Oxidative stress in the body is believed to contribute to chronic diseases like liver disease, cardiovascular disease, certain types of cancer, and neurodegenerative diseases.³³⁻³⁵ Antioxidant-rich foods have shown positive effects on cardiovascular health, cancer prevention, and cognitive function.³⁶⁻³⁸ By neutralizing free radicals and reducing oxidative stress, a diet high in antioxidants may help prevent the progression of chronic diseases.³⁹ In line with these findings, our study revealed a negative correlation between antioxidant intake and the prevalence of CRDs. Specifically, individuals with higher total dietary antioxidant intake exhibited a lower prevalence of emphysema and chronic bronchitis. These results lend support to the notion that antioxidants may play a protective role in the development of CRDs.

Antioxidants play a crucial role in maintaining the delicate balance between harmful ROS and protective antioxidants in the body.⁴⁰ Research indicates that increasing antioxidant consumption in the diet can reduce the incidence of respiratory disorders.⁴¹ Notably, vitamins A, C, and E are known for their potent antioxidant properties and immune-enhancing effects.⁴² These vitamins play a vital role in safeguarding the respiratory tract from inflammation, which is a common underlying factor in diseases like asthma and COPD.⁴³ Additionally, zinc and selenium, as trace minerals, act as cofactors for antioxidant enzymes in the body. They contribute to normal immune system functioning by reducing inflammation and promoting tissue healing in the respiratory system.^{44,45} Deficiencies in these minerals are associated with impaired respiratory function and heightened susceptibility to respiratory infections.⁴⁶

Carotenoids, found in various foods, are pigments with antioxidant properties. They have demonstrated protective effects on the respiratory system, including a reduced risk of asthma and COPD.⁴⁷ Overall, our findings suggest that higher CDAI scores (indicating multiple antioxidant consumption) are associated with a decreased likelihood of respiratory diseases when compared to single antioxidant intake. It is crucial to highlight that the link between CDAI and respiratory health is complex and influenced by other factors such as genetics, lifestyle, and environmental exposures.

The protective effects of antioxidants can be attributed to several mechanisms. Firstly, oxidative stress has been linked to the development of CRDs, leading to airway inflammation and tissue damage.⁴⁸ Antioxidants work to neutralize free radicals and protect cells and tissues from oxidative damage.⁴⁹ Therefore, a higher dietary antioxidant intake may reduce oxidative stress, alleviating symptoms and halting the progression of CRDs. Secondly, antioxidants have been found to influence immunological activity, a critical characteristic of many CRDs.^{50,51} Antioxidants, such as vitamin C and vitamin E, possess immunomodulatory properties that promote a balanced immune response, modulate the production of pro-inflammatory mediators, and reduce allergic reactions.^{52,53} This immunomodulatory action could help explain the apparent protective relationship between dietary antioxidant intake and CRDs. Furthermore, antioxidants have been linked to enhanced lung function.⁵⁴ Oxidative stress-induced lung tissue damage can lead to impaired respiratory function.⁵⁵ Antioxidant-rich diets, particularly those high in fruits and vegetables, have been associated with improved lung function indicators.⁵⁶ These findings suggest that increasing dietary antioxidant consumption may promote lung health and lower the prevalence of CRDs.

Our study identified several key risk factors associated with CRDs, including older age (≥ 60 years), female gender, non-Hispanic white ethnicity, current smoking status, obesity, and physical inactivity. These findings underscore the multifactorial nature of CRDs and the importance of addressing the complex interplay of factors that

contribute to their development. Inflammation is a central player in the pathogenesis of CRDs. Current smokers and obese individuals are known to have higher rates of inflammation due to increased production of pro-inflammatory cytokines such as TNF α and IL-6 from adipose tissue, as well as a lower production of anti-inflammatory cytokines.⁵⁷ This heightened inflammatory state can contribute to the progression of CRDs. It is noteworthy that our results also indicated a higher prevalence of hypertension and diabetes among individuals with CRDs. Both of these conditions are closely linked with inflammation, and in particular, weight status plays a pivotal role in this relationship. Obesity is recognized as a state of chronic low-grade inflammation, which can exacerbate the risk of hypertension and diabetes.⁵⁷ Furthermore, our findings revealed that CRD patients were more likely to be over the age of 60 years and female. This aligns with the observation that postmenopausal women, experiencing lower estrogen levels, face an increased risk of several chronic diseases. Changes in fat distribution in the body during menopause contribute to a pro-inflammatory environment,⁵⁸ further underscoring the role of inflammation in the pathogenesis of CRDs.

The study's strength is that it uses data from an extensive number of participants from the nine cycles of the 2001-2018 NHANES, which increases the confidence of the study's findings. Second, compared with previous metrics, the CDAI is a novel approach to assessing dietary antioxidant intake that takes into account the intake of key antioxidants, including vitamins A, C, and E, and zinc, selenium, and carotenoids. Third, we not only explored the effects of multiple confounders, but also assessed the association between CDAI and the prevalence of CRDs through stratified analyses.

However, this study has several drawbacks. First, this is a cross-sectional study, which does not allow for causal inference of correlations. Second, CRDs include not only asthma, emphysema and chronic bronchitis but also other diseases, so the conclusions drawn from this study are limited and need to be validated by including more types of CRDs. Third, the intakes of various dietary nutrients were derived from the average of 2 24-h dietary reviews, so they may not be representative of daily dietary changes. Additionally, dietary intakes may vary

among individuals due to personal preferences and cultural factors, leading to potential confounding effects.

CONCLUSION

CDAI is negatively associated with the prevalence of CRDs (especially emphysema and chronic bronchitis) in the adult population. These findings highlight a potential benefit to diets high in antioxidant-rich foods with regards to lung function and onset of CRD. Further studies need to be done to determine whether higher dietary intake of antioxidant-rich foods is in fact protective especially in populations at higher risk of developing CRDs (ie, obese, postmenopausal women, or physically inactive individuals). Further studies on the underlying mechanisms are necessary.

Abbreviations

CRDs, Chronic respiratory diseases; NHANES, National Health and Nutrition Examination Survey; CDAI, composite dietary antioxidant index; RCS, restricted cubic spline; ROS, reactive oxygen species; COPD, chronic obstructive pulmonary disease; NCHS, National Center for Health Statistics; PIR, poverty income ratio; MET, metabolic equivalents; SEs, standard errors; IQRs, interquartile ranges.

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Data availability

NHANES data described in this manuscript are available at <https://www.cdc.gov/nchs/nhanes/>.

Author contributions

Shidong Wang: Conceptualization, Methodology, Software, Formal analysis, Writing-original draft, Visualization. Hong Teng: Writing-review and editing. Lin Zhang: Writing-review and editing. Liang Wu: Conceptualization, Methodology, Project administration, Writing-review and editing, Supervision.

Ethics approval and consent to participant

All participants provided written informed consent and study procedures were approved by the National Center for Health Statistics Research Ethics Review Board (Protocol Number: Protocol #98-12, Protocol #2005-06, Protocol #2011-17, and Protocol #2018-01).

Consent to participate

The manuscript is approved by all authors for publication.

Declaration of competing interest

The authors have no competing interests to declare.

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Appendix A. Supplementary data

Supplementary data to this article can be found online at <https://doi.org/10.1016/j.waojou.2023.100851>.

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