Association between total dietary antioxidant capacity and food groups and incidence of depression in a cohort of Brazilian graduates (CUME Project)

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(Submitted 27 September 2022 – Final revision received 16 December 2022 – Accepted 16 January 2023 – First published online 1 February 2023)

Abstract

This study aims to evaluate the association between Dietary Total Antioxidant Capacity (dTAC) and Total Antioxidant Capacity of food groups (fgTAC) with the incidence of depression in Brazilian graduates participating in the Cohort of Universities of Minas Gerais (CUME Study). The sample consisted of 2572 participants without a medical diagnosis of depression at baseline who responded to at least one follow-up questionnaire from the CUME Project. The Ferric Reducing Antioxidant Power assay was used to determine dTAC. Incidence of depression was estimated by self-reported medical diagnosis of depression during the years of cohort follow-up. Cox regression models were used to relate dTAC and fgTAC to the incidence of depression. The mean follow-up time was 2.96 (1.00) years, and 246 cases of depression were observed (32-3/1000 person-years). The mean dTAC was 11.03 (4.84) mmol/d. We found no associations between higher dTAC and lower risk of developing depression after adjusting for possible confounders. The incidence of depression was inversely associated with fgTAC of the beans and lentils group (hazard ratio (HR): 0.61; 95 % CI 0.41, 0.90). The fgTAC of the junk food group was positively associated with higher incidence of depression after all adjustments (HR: 1.57; 95 % CI 1.08, 2.26). Our findings do not support an association between dTAC and the incidence of depression in a highly educated Brazilian population. However, associations of fgTAC show the importance of analysing the food matrix in which these antioxidants are inserted. We highlight the need for more prospective studies with different nationalities to confirm these results.

Key words: Dietary Total Antioxidant Capacity: Depression: Epidemiology: Oxidative stress

Depression is a chronic mental illness with great worldwide prevalence⁽¹⁾. Data indicate that 3.8% of the world population is affected by depression, affecting about 5% of the adult population and 5.7% of the population over 60 years old⁽¹⁾. The main characteristics of depressive disorder are depressed mood, feelings of guilt or low self-esteem, changes in sleep and appetite, lack of disposition and poor concentration. In many cases, these symptoms can be accompanied by feelings of anxiety⁽²⁾. In moderate or severe intensities, depression can greatly affect an individual's daily life in work, studies and family relationships, and in its most severe forms, it can lead to suicide^(1,2).

In this sense, much has been investigated about the pathogenesis of depression, and oxidative stress has stood out among the risk factors^(3,4). Oxidative stress can be characterised as an imbalance between the antioxidant defences of the organism and the presence of free radicals, a pathogenic process related to cell injury and death^(3–5). It is noteworthy that the brain, compared with other organs, is highly vulnerable to oxidative stress due to its high metabolism^(5,6). Thus, redox imbalance may be related to depression through mechanisms such as inflammation and neurodegeneration, impairing neuronal and neurotransmitter function^(3,4,7).

Due to the need for new approaches to treat and prevent depression, antioxidants such as vitamins C, E and Zn have been associated with improvements in neurocognitive function, bringing therapeutic benefits to depression⁽⁸⁾. A recent cross-sectional study with 14 737 individuals participating in the Brazilian Longitudinal Study of Adult Health (ELSA) reported,



Abbreviations: dTAC, Dietary Total Antioxidant Capacity; fgTAC, Total Antioxidant Capacity of food groups; FRAP, Ferric Reducing Antioxidant Power.

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for Brazilian women, an inverse association of the consumption of Zn, Se, vitamin A and C with depression⁽⁹⁾. Inverse associations of Zn and Se consumption with depression were also found for 14 834 adults participating in the National Health and Nutrition Examination Survey $2009-2014^{(10)}$.

Despite the results obtained so far between nutrient intake and the occurrence of depression, the isolated assessment of antioxidants may not be as effective as the assessment of the interaction of different dietary antioxidants⁽¹¹⁾. Therefore, the Dietary Total Antioxidant Capacity (dTAC), an index capable of measuring the global content of dietary antioxidants⁽¹²⁾, has proved to be a helpful tool for investigating the interaction between dietary antioxidants and health outcomes⁽¹³⁻¹⁵⁾. However, investigations into the relationship between dTAC and depression are still limited, mostly being cross-sectional studies with the Iranian population⁽¹⁶⁻¹⁹⁾. Unfortunately, as far as we know, no prospective study has investigated the association between dTAC and depression in Brazilians, nor has it analysed this relationship through the Total Antioxidant Capacity of food groups (fgTAC)⁽¹⁶⁾. Thus, the present study aimed to assess the association between dTAC and fgTAC with the incidence of depression in Brazilian graduates participating in the Cohort of Universities of Minas Gerais (CUME Study).

Methodology

Cohort of Universities of Minas Gerais (CUME Study)

The CUME Study is an open, prospective cohort conducted with alumni from universities located in the state of Minas Gerais (Brazil), whose main objective is to assess the impact of the Brazilian dietary pattern, specific diet factors and the nutritional transition in the incidence of chronic non-communicable diseases, as previously detailed⁽²⁰⁾.

The CUME Study questionnaires were developed on the Alchemer (www.alchemer.com) online interface by specialists. The team conducted pilot studies with the baseline questionnaire to assess its applicability⁽²⁰⁾. The CUME Study began in 2016 with the application of the baseline questionnaire Q_0, and its recruitment has been periodic since then. Invitations to participate were sent to all volunteers who had emails available. Thus, every 2 years, the participants are invited to answer follow-up questionnaires (Q_2, Q_4...Q_n) to update their information about lifestyle, the emergence of new diseases and changes in dietary patterns, among others, while new potential participants are recruited and invited to answer the baseline cohort questionnaire (Q_0). Characteristics related to project design and recruitment of the first volunteers were described in a previous study⁽²⁰⁾.

The questionnaires are answered in a virtual environment of the CUME Study. Q_0 is divided into two phases: the first phase with questions about socio-demographic characteristics, lifestyle, biochemical markers (TAG concentrations, total cholesterol, HDL-cholesterol, LDL-cholesterol, blood glucose concentration) and related to the individual's health outcomes; the second phase that is answered after 1 week contains a FFQ and questions related to dietary practices. On the other hand, Q_2 is composed of questions related to changes in lifestyle, eating habits and health conditions. Finally, the follow-up questionnaire Q_4 has questions related to socio-demographic characteristics, lifestyle, biochemical markers, health outcomes and insomnia severity.

This study was conducted according to the guidelines laid down in the Declaration of Helsinki, and all procedures involving human subjects were approved by the Human Research Ethics Committees of all participating institutions: Federal University of Minas Gerais (CAAE registration number: 07223812.3.3001.5153); Federal University of Viçosa (CAAE registration number: 4483415.5.1001.5149); Federal University of Ouro Preto (CAAE registration number: 44483415.5.2003.5150); Federal University of Lavras (CAAE registration number: 44483415.5. 2002.5148); Federal University of Juiz de Fora (CAAE registration number: 4483415.5.5133); Federal University of Vale do Jequitinhonha e Mucuri (CAAE registration number: 44483415.5. 2005.5103) and Federal University of Alfenas (CAAE registration number: 4.501.344). Written informed consent was obtained from all subjects⁽²⁰⁾.

Data collection

Information from the Q_0 questionnaires for the years 2016, 2018 and 2020 composed the baseline of the present study. We used data from the 2-year follow-up Q_2 and the 4-year follow-up Q_4 to investigate the incidence of the depression outcome.

The first collection related to the 2-year follow-up occurred in 2018, answered by participants who started the baseline (Q_0) in 2016. The second collection, associated with Q_2, occurred in 2020 when we invited participants who completed the Q_0 follow-up questionnaire in 2018. Finally, the collection related to the Q_4 follow-up questionnaire occurred in 2020. Participants who answered Q_2 in 2018 were invited to fill it out.

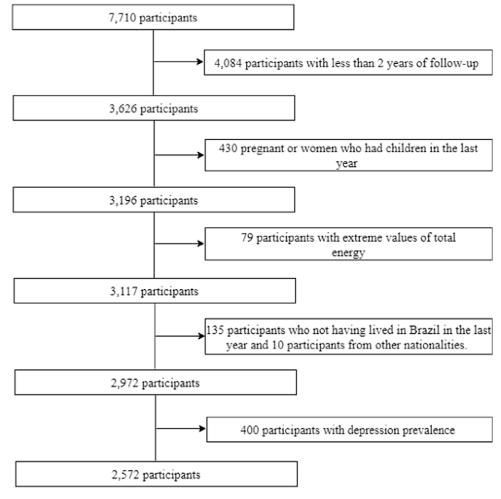
Study population

The CUME Study has responses from 7710 participants. For the present study, of the total number of respondents, we excluded 4084 participants who had not accomplished at least the 2-year follow-up. We also excluded women who were pregnant or had been pregnant in the last year (n 430), participants with energy consumption < 2092 or > 25104 kJ per day (< 500 kcal or > 6000 kcal) (n 79)⁽²¹⁾, participants who reported not having lived in Brazil in the last year (n 135) and foreigners living in Brazil (n 10) and participants who reported a diagnosis of depression at baseline (n 400). The final sample consisted of 2572 participants who answered at least one follow-up questionnaire (Fig. 1).

Outcome variable: incidence of depression

We considered incident cases of depression in those participants who were disease-free at the beginning of the follow-up and were classified as having the disease during Q_2 or Q_4 follow-up. For it, we considered incident depression when participants answered yes to the following question: 'Since the previous questionnaire, were you clinically diagnosed with depression for the first time?'. Individuals who reported using antidepressants but did not confirm the diagnosis of depression

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Fig. 1. Flow chart of participant selection.

were not included due to the possibility of the therapeutic use of such drugs in diseases other than depression. The reliability between medical diagnosis and self-report depression in this cohort was validated in a subsample of participants showing good agreement (81.0%) with a κ value of 0.62⁽²²⁾.

Food consumption and estimation of Dietary Total Antioxidant Capacity

We assessed habitual food consumption using FFQ, previously validated for the study population⁽²³⁾, which includes 144 food items presented in the following groups: dairy products, meat and fish, cereals and legumes, fruits, vegetables, fats, and oils, beverages and other foods. Each participant reported the frequency of consumption of a particular food (daily, weekly, monthly or yearly), the number of times they consumed it (0–9 or more times) and the portion size. To facilitate filling in the portion sizes of food items and obtain the most reliable information possible, participants had access, at the time of answering the FFQ, to images of portions of food and serving utensils from the photo album with ninety-six pictures prepared by the CUME Study team⁽²⁰⁾. The consumption of macro and micronutrients was calculated using primarily data from the

Table of Nutritional Composition of Foods consumed in Brazil⁽²⁴⁾ and, in the absence of information in this table, we consulted the Brazilian Table of Food Composition⁽²⁵⁾ and the USDA National Nutrient Database⁽²⁶⁾.

We used the values from the Ferric Reducing Antioxidant Power (FRAP) assay to estimate dTAC, which measures the antioxidant capacity of food in the presence of Fe. We consulted previously published databases to obtain the FRAP values^(12,27). Thus, the dTAC of each food item resulted from the multiplication between the amount in grams of food consumed and its corresponding FRAP value in mmol per gram of food (mmol/g).

We used the following criteria to assign the FRAP value to a portion of food: when there was more than one FRAP analysis value for the same food, we considered the mean value; in the absence of a FRAP value for a particular food, when possible, we used the value of a similar food from the same botanical group, or the same food in a different way of preparation. We did not assign FRAP values to foods where there was no record of this value, and it was not possible to estimate for foods from similar botanical groups or different methods of preparation

A total of 133 food items were covered with FRAP values. We summed all FRAP values of foods reported in the FFQ to estimate the total dTAC of each participant. In order to perform a

https://doi.org/10.1017/S0007114523000181 Published online by Cambridge University Press

sensitivity analysis, we also calculated dTAC values, excluding coffee values, as this is a beverage that greatly contributes to dTAC values in our population⁽¹³⁾. We also calculated fgTAC values according to food groups (fruits, vegetables, beans and lentils, oilseeds, dairy products, meats and eggs, pasta, breads and cereals, oils and fats, junk food, natural juices, teas and coffees, artificial juices and sodas, and alcoholic beverages) (online Supplementary Table 1).

We adjusted all food consumption variables for daily energetic intake using the residual method⁽²⁸⁾, including dTAC.

Covariates

We obtained the other variables from self-reported information in the baseline questionnaire O 0, including socio-demographic variables (sex, skin colour, marital status, professional status), use of vitamin supplements and smoking habits (non-smoker, smoker or ex-smoker). The frequency of heavy episodic consumption of alcoholic beverages (1-2 d/month, 3-4 d/month and 5 or more days/month) was also a variable included in this study, with heavy episodic consumption considered as 4 or more doses of alcoholic beverage on a single occasion for women and 5 or more doses of alcoholic beverage on a single occasion for men⁽²⁹⁾. We also assessed physical activity using a list of twentyfour activities expressed in minutes per week. Participants with ≥ 150 min/week of moderate-intensity activity or ≥ 75 min/week of vigorous-intensity activity were considered active; participants with < 150 min/week of moderate-intensity activity or <75 min/week of vigorous-intensity activity were classified as insufficiently active and those who reported no leisure-time physical activity were classified as inactive⁽³⁰⁾. We obtained the BMI from the self-reported weight and height in the baseline questionnaire, dividing weight (kg) by height (m) squared. BMI values ≥ 30 kg/m² were classified as obesity^(31,32). The BMI based on self-reported weight and height was also validated, in a previous study, for the population participating in the CUME Project, indicating excellent agreement with the measured intra-class correlation coefficient 0.989⁽³³⁾.

Statistical analysis

We performed the analyses with Stata SE 15.0. A two-tailed *P*-value less than 0.05 was considered statistically significant.

We described socio-demographic, lifestyle, health and food consumption characteristics in absolute or relative frequency or as mean and standard deviation according to the dTAC quartiles baseline. We calculated the *P*-value using Pearson's chi-squared tests for categorical variables and ANOVA for continuous variables to compare the categories.

The follow-up time was calculated in person-years for each participant: difference between the date of completion of the follow-up questionnaire in which depression was diagnosed and the date of completion of the baseline questionnaire. In this sense, we created Cox regression models to assess the association between dTAC in quartiles and the incidence of depression. We used the lowest quartile as the reference category to compare the incidence of depression among the dTAC categories (exposure variable). As a guide for selecting the covariates included in the analyses, we constructed a directed acyclic graph (DAG) using the DAGitty program. DAG are a strategy that helps identify a minimum set of confounding covariates in the analysis of causal relationships, helping to estimate less biased measures of effect^(34,35) (online Supplementary Fig. 1). Thus, we adjusted the final model for potential confounders such as sex, age (continuous), smoking habit (non-smoker/smoker/ex-smoker), frequency of heavy episodic alcohol consumption (never/from 1 to 2 days per month/from 3 to 4 days per month/5 or more days per month), marital status (single/married or stable union/separated or divorced or widowed), skin colour (white/non-white), professional status, physical activity (inactive/insufficiently active/active), use of vitamin supplements (yes/no), BMI (continuous) and energetic intake (kcal/d) and vitamin D consumption (μ g/d). Linear trends were tested using the median values of each quartile of the exposure variable ordered in Cox regression models. We also performed a sensitivity analysis excluding coffee items from the dTAC computation. In addition, we analysed the relationship between fgTAC and the incidence of depression (fruits, vegetables, beans and lentils, oilseeds, dairy products, meat and eggs, breads, pasta and cereals, oils and fats, 'junk food', natural juices, teas and coffees, artificial juices and soda and alcoholic beverages), considering that antioxidants present in diet may be inserted in different proportions in food matrices (15,36).

Results

During the average time of 2.96 (1.00) years of monitoring of the present study, 246 new cases of depression (32.3/1000 personyears) were observed and the mean age of the participants was 36.09 (SD 9.63). The mean dTAC was 11.03 (4.84 mmol/d).

Participants included in the highest quartile for dTAC (> 13·32 mmol/d) are mostly older, married or in a stable union, employed workers, smokers and physically active, in addition to having a higher frequency of heavy episodic alcohol consumption (Table 1). Regarding food consumption, individuals belonging to the fourth quartile of dTAC, when compared with the first quartile (< 7·92 mmol/d), had a higher intake of carbohydrates, *n*-3, alcohol, vitamins A, E, C and B₉, Mg, fibre, as well as higher consumption of fruits and vegetables (Table 2).

There was no association between dTAC and incidence of depression (Table 3), regardless of adjustment for confounding variables. The results of the analysis of dTAC without coffee and incidence of depression remained similar. When we assessed the associations between fgTAC and the incidence of depression (Tables 4 and 5), we observed a lower incidence of depression according to the quartiles of fgTAC of natural juices in the model adjusted by age and sex (hazard ratio: 0.70; 95 % CI 0.49, 0.99). Still, the significance of this association was lost after the total adjustment of the model. On the other hand, the TAC of the beans and lentils was inversely associated with the incidence of depression in our cohort (hazard ratio: 0.61; 95 % CI 0.41, 0.90) (Table 4). Interestingly, a higher fgTAC from the junk food group was positively associated with a higher incidence of depression among participants after all adjustments (hazard ratio: 1.57; 95 % CI 1.08, 2.26).

https://doi.org/10.1017/S0007114523000181 Published online by Cambridge University Press

Table 1. Baseline socio-demographic and health characteristics according to energy-adjusted dTAC (mmol/d) quartiles, CUME Project (*n* 2572) (Mean values and standard deviations; numbers and percentages)

		Q1		Q2		Q3		Q4	
Mean sd	6·02 1·80		9·09 0·67			1.69 0.85	17·33 4·49		
00	n	%	n	%	n	%	n	%	P
Age									
<40 years	517	80.4	476	74·0	421	65.5	393	61.1	<0.001
40–59 years	115	17.9	152	23.6	200	31.1	231	35.9	
\geq 60 years	11	1.7	15	2.4	22	3.4	19	3.0	
Sex									
Male	251	39.0	214	33.3	219	34.1	252	39.2	0.040
Female	392	61·0	429	66.7	424	65.9	391	60.8	
Skin colour									
White	406	63·1	403	62·7	418	65·0	434	67.5	0.257
Non-white	237	36.9	240	37.3	225	35.0	209	32.5	
Marital status									
Single	370	57.5	315	49.0	281	43.7	277	43.1	
Married/stable union	250	38.9	296	46.0	322	50.1	313	48.7	<0.00
Separated/divorced/widower/other	23	3.6	32	5.0	40	6.2	53	8.2	
Professional situation									
Employee	445	69.2	506	78.7	495	77.0	508	79.0	
Student	126	19.6	87	13.5	92	14.3	86	13.4	<0.001
Retired	9	1.4	9	1.4	17	2.6	18	2.8	
Unemployed	63	9.8	41	6.4	39	6.1	31	4.8	
Smoking status									
No	565	87.9	533	82.9	502	78.1	475	73.9	
Past	38	5.9	69	10.7	86	13.4	91	14.2	<0.00
Current	40	6.2	41	6.4	55	8.6	77	12.0	
Binge frequency									
None	401	62.4	402	62.5	370	57.5	327	50.9	
1–2 times a month	124	19.3	130	20.2	148	23.0	147	22.9	<0.00
3–4 times a month	75	11.7	75	11.7	69	10.7	89	13.8	
5 or more times a month	43	6.7	36	5.6	56	8.7	80	12.4	
Physical activity									
Inactive	204	31,7	156	24,3	128	19,9	110	17,1	
Insufficiently active	134	20,8	141	21,9	130	20,2	118	18,4	<0.001
Active	305	47,4	346	53,8	385	59,9	415	64,5	
Use of supplements		*		*		*			
Yes	149	23.2	157	24.4	181	28.1	182	28.3	0.078
No	494	76.8	486	75.6	462	71.9	461	71.7	
Obesity									
No	561	87·2	574	89.3	582	90.5	579	90.0	0.242
Yes	82	12.8	69	10.7	61	9.5	64	10.0	

dTAC, Dietary Total Antioxidant Capacity.

P values according to trend chi-squared test

Discussion

In the present study, we found no association between total dTAC and incidence of depression. Still, fgTAC from specific food groups, such as natural juice, beans and lentils, and junk foods, showed associations. As far as we know, this is the first prospective study to investigate the association between dTAC and fgTAC by different food groups with the incidence of depression in Brazilian graduates.

When analysing the fgTAC, we observed an inverse association between the fgTAC of the beans and lentils group with the incidence of depression. A study from the National Health Survey in Brazil with 46 785 adults showed an inverse association between bean consumption and depression⁽³⁷⁾. Bean consumption was also inversely associated with mental disorders in a study with 712 Brazilian pregnant women⁽³⁸⁾. It is noteworthy that beans, besides being sources of antioxidants such as polyphenols, are sources of B vitamins and minerals such as Fe, K and Mg, in addition to dietary fibre^(39,40). In a previous study, beans proved to be an important contributor to folate consumption in part of the CUME Project baseline population⁽⁴¹⁾. In fact, in addition to antioxidants, folate consumption has been inversely associated with depression^(42,43). Another point worth mentioning is the fact that the regular consumption of beans may be related to a higher quality dietary pattern, characterised by a diversity of in natura and minimally processed foods, while low intake may be associated with an increase in the consumption of ultra-processed foods^(40,44,45). We emphasise that the dietary pattern rich in ultra-processed foods is positively related to the incidence of depression^(37,46).

We observed that higher fgTAC of junk food was positively associated with higher incidence of depression in our cohort. This fact raises the question of the importance of the food matrix in which antioxidants are inserted. Although some ultra-processed foods have vitamins and minerals with antioxidant

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Table 2. Baseline dietary intake according to the energy-adjusted dTAC (mmol/d) quartiles, CUME Project (n 2572) (Mean values and standard deviations)

	C	21	C	2	(23	(Q4		
Mean SD		02 80		09 67		·69 ·85		7.33 .49		
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	P*	
Total energy intake, kcal/d	2662.50	1027.93 ^a	2091.645	761.19 ^b	2200.05	793·04 ^b	2504.59	986.68°	<0.001	
Carbohydrates, g/d	241.54	67.05ª	259.92	44.43 ^b	265.17	54.59 ^{b,c}	272.72	69.11°	<0.001	
Protein, g/d	113.17	36.38 ^a	105.92	23·26 ^b	103.60	28.00 ^b	99.39	30.82 ^c	<0.001	
Lipids, g/d	101.63	23.60 ^a	96.14	16·13 ^b	93.93	19.02 ^{b,c}	90.99	24.36 ^c	<0.001	
SFA, g/d	36.09	10·28ª	33.58	6.80 ^b	32.51	8.94 ^{b,c}	30.93	9.62 ^c	<0.001	
MUFA, g/d	36.15	10.46 ^a	35.07	7.52 ^{a,b}	34.48	9.25 ^{b,c}	33.28	11.11°	<0.001	
PUFA, g/d	18.36	6.99ª	19.24	5.08 ^a	19.33	5.40 ^a	19.15	7.38 ^a	0.021	
TRANS, g/d	1.41	1.27ª	1.22	0.73 ^b	1.10	0.70 ^b	0.93	0.80 ^c	<0.001	
n-3 fatty acids, g/d	2.09	0.73ª	2.30	0.55 ^{b,c}	2.38	0.55 ^c	2.60	0.87 ^d	<0.001	
Alcohol intake, g/d	4.43	9.89 ^a	5.14	6.46 ^{a,b}	6.31	7.97 ^b	8.17	11.92 ^c	<0.001	
Vitamin C, mg/d	161.34	134.79 ^a	236.67	151.71 ^b	277.50	196.60 ^c	335.43	286·27 ^d	<0.001	
Vitamin A, μg/d	761·89	324·25 ^a	835.82	291.15 ^b	869.10	280.26 ^{b,c}	912·19	546·31°	<0.001	
Vitamin E, mg/d	7.39	3.16ª	8.51	2.64 ^b	9.11	3.16°	9.95	4.75 ^d	<0.001	
Folic acid, µg/d	450.31	179.77 ^a	497.34	113·24 ^b	497.45	124.52 ^b	509.58	142·11 ^b	<0.001	
Vitamin B ₁₂ , µg/dc	4.55	3.13ª	4.22	1⋅81 ^{b,a}	4.05	1.65 ^b	3.84	3.11 ^b	<0.001	
Vitamin D, µg/d	4.22	3.57ª	4.20	2.82 ^a	4.04	2.57 ^{a,b}	3.73	2.67 ^b	0.008	
Mg, mg/d	324.88	81.93 ^a	366.15	72.53 ^b	391.66	77.50 ^c	424.95	98.65 ^d	<0.001	
Fibre, g/d	21.06	9.70 ^a	25.88	7.07 ^b	28.10	8.59 ^c	30.80	12.22 ^d	<0.001	
Fruit, g/d	267.45	248.88 ^a	414.15	218·94 ^b	491.77	290.02 ^c	598·97	441.48 ^d	<0.001	
Vegetables, g/d	188.75	142.55 ^a	222.26	121.12 ^b	245.91	132·15 ^{c,d}	262.90	178·20 ^d	<0.001	
Legumes, g/d	93.46	120.75 ^a	80.65	83·50 ^{a,b}	75.78	83·46 ^b	80.15	84·14 ^{a,b}	0.005	

Data expressed as mean (standard deviation).

* P values by ANOVA test. Different letters show statistically significant differences between groups according to Bonferroni's post hoc test.

Table 3. Hazard ratios and 95 % CI of depression incidence according to dTAC and dTAC without coffee, CUME Project (n 2572) (Mean values and standard deviations)

				Total dTA	C			
	Q1		Q2		Q3		Q4	P _{for trend}
Mean	6.02		9.09		11.69		17.33	
SD	1.80		0.67		0.85		4.49	
Crude	1.00	0.98	0.68, 1.41	1.15	0.81, 1.63	1.07	0.75, 1.52	0.591
Model 1*	1.00	0.98	0.68, 1.41	1.16	0.81, 1.67	1.10	0.77, 1.59	0.460
Model 2†	1.00	0.97	0.67, 1.40	1.14	0.79, 1.64	1.09	0.75, 1.58	0.521
Model 3‡	1.00	1.10	0.75, 1.62	1.29	0.88, 1.87	1.18	0.81, 1.72	0.342
				dTAC without	coffee			

	Q1		Q2		Q3		Q4	P _{for trend}
Mean	4.08		6.29		7.76		11.79	
SD	1.42		0.39		0.49		3.85	
Crude	1.00	1.16	0.81, 1.65	0.94	0.65, 1.36	1.23	0.86, 1.76	0.370
Model 1*	1.00	1.12	0.78, 1.59	0.88	0.61, 1.29	1.15	0.79, 1.66	0.605
Model 2†	1.00	1.11	0.78, 1.59	0.88	0.60, 1.28	1.16	0.80, 1.68	0.562
Model 3‡	1.00	1.29	0.89, 1.87	1.02	0.69, 1.50	1.32	0.91, 1.91	0.246

dTAC, Dietary Total Antioxidant Capacity.

Adjusted - Sex and age.

Adjusted – Model 1 + smoking status (never, current, former), alcohol consumption (BINGE frequency) and vitamin D consumption (mcg).
‡ Adjusted – Model 2 + marital status (single/married or stable union/separated or divorced or widowed), skin colour (white and not white), physical activity (inactive/insufficiently) active/active), use of supplements (yes or no), energy intake (continuous, kcal/d), baseline BMI (continuous kg/m²), professional situation.

potential added to their composition to increase their shelf life, these are generally rich in simple sugars, fats, flavourings and preservatives that can contribute to a pro-oxidant and inflammatory state, closely related to depression⁽⁴⁷⁻⁴⁹⁾. In addition, a diet rich in fast food can contain a lower amount of vitamins and minerals than in natura or minimally processed foods. The deficient consumption of several nutrients is related to depression^(50,51).

We did not observe any association between total dTAC and incidence of depression. These findings agree with a prospective study with 911 Japanese workers, with no associations between dTAC and incidence of depressive symptoms after 3 years of

Table 4. Hazard ratios and 95 % CI of depression incidence according to fgTAC from food groups, CUME Project (n 2572) (Mean values and standard deviations)

				fgT	AC from fruits			
	Q1		Q2		Q3		Q4	P _{for tren}
Mean	0.49		1.76		2.68		5.39	
SD	0.79		0.25		0.30			
Crude	1.00	1.14	0.80, 1.62	0.97	0.67, 1.40	1.13		0.668
Model 1*	1.00	1.06	0.74, 1.51	0.87	0.60, 1.26	1.03	,	0.962
Model 2†	1.00	1.06	0.74, 1.51	0.87	0.60, 1.20	1.04		0.996
Model 3‡	1.00	1.26	0.87, 1.83	1.03	0.70, 1.53	1.23	0.84, 1.80	0.473
				fgTAC	from vegetables			
	Q1		Q2		Q3		Q4	P _{for trend}
Mean	0.26		0.53		0.72		1.29	
SD	0.15		0.05		0.06		0.60	
Crude	1.00	0.84	0.59, 1.18	0.96	0.68, 1.36	0.75	0.53, 1.08	0.187
Model 1*	1.00	0.79	0.56, 1.11	0.89	0.63, 1.25	0.72	0.50, 1.14	0.135
Model 2†	1.00	0.78	0.55, 1.11	0.88	0.62, 1.25	0.71	0.49, 1.03	0.121
Model 3‡	1.00	0.86	0.60, 1.23	0.96	0.67, 1.37	0.77	0.53, 1.11	0.216
				fgTAC fro	m beans and lentils		3·32 0·79, 1·61 0·71, 1·49 0·72, 1·50 0·84, 1·80 Q4 1·29 0·60 0·53, 1·08 0·50, 1·14 0·49, 1·03	
	Q1		Q2		Q3		Q4	P _{for trend}
Mean	0.02		0.15		0.26			
SD	0.07		0.03		0.03			
Crude	1.00	1.09	0.79, 1.49	0.63	0.44, 0.90	0.56		0.001
Model 1*	1.00	1.06	0.77, 1.45	0.63	0.44, 0.90	0.59		0.001
Model 2†	1.00	1.03	0.75, 1.43	0.61	0.42, 0.88	0.57	0.38, 0.83	0.001
Model 3‡	1.00	1.14	0.81, 1.60	0.68	0.46, 1.00	0.61	0.41 - 0.90	0.002
				fgTA	C from oil seeds			
	Q1		Q2		Q3		Q4	P _{for trend}
Mean ^{SD}	0·01 0·24		0·19 0·07		0·43 0·08			
Crude	1.00	0.71	0.49, 1.03	0.96	0.67–1.36	1.08		0.216
Model 1*	1.00	0.65	0.44, 0.94	0.90	0.59, 1.21	0.98		0.210
Model 2†	1.00	0.65	0.45, 0.95	0.85	0.60, 1.23	0.99		0.351
Model 3‡	1.00	0.05	0.40, 0.95	1.05	0.69, 1.60	1.13		0.331
				fgTAC f	rom dairy products		0.60 0.35 0.38, 0.81 0.40, 0.86 7 0.38, 0.83 0.41 - 0.90 0.41 - 0.90 0.41 0.41 - 0.90 0.41 - 0.90 0.	
	Q1		Q2	-	Q3		Q4	P _{for trend}
Mean	0.03		0.09		0.15		0.29	
SD	0.03		0.01		0.02		0.11	
Crude	1.00	0.92	0.63, 1.34	1.03	0.72, 1.48	1.19	0.83, 1.70	0.215
Model 1*	1.00	0.90	0.62, 1.30	0.99	0.69, 1.42	1.15	0.80, 1.64	0.296
Model 2†	1.00	0.92	0.63, 1.34	1.02	0.70, 1.48	1.20		0.217
Model 3‡	1.00	0.96	0.66, 1.39	1.05	0.73, 1.52	1.22		0.204
				fgTAC fr	om meat and eggs			
	Q1		Q2		Q3		Q4	P _{for trend}
Mean	0.06		0.15		0.21		0.39	
SD	0.06		0.01		0.09		0.18	
Crude	1.00	0.94	0.66, 1.34	0.86	0.60, 1.24	1.06	0.75, 1.49	0.777
Model 1*	1.00	0.88	0.62, 1.25	0.82	0.57, 1.18	1.00	0.71, 1.42	0.948
Model 2†	1.00	0.90	0.63, 1.27	0.84	0.58, 1.21	1.01	0.69, 1.47	0.929
Model 3‡	1.00	0.98	0.67, 1.42	0.92	0.62, 1.36	1.03	0.70, 1.52	0.872
				fgTAC from b	read, pasta and cerea	als		
	Q1		Q2		Q3		Q4	P _{for trend}
 Mean	0.21		0.40		0.56		0.97	

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Table 4. (Continued)

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				fgTAC from b	ead, pasta and cerea	als		
	Q1		Q2		Q3		Q4	P _{for tren}
Crude	1.00	0.81	0.56, 1.16	0.87	0.62, 1.24	0.89	0.63, 1.26 0.61, 1.24 0.61, 1.24 0.65, 1.31 Q4 0.30 0.14 0.72, 1.46 0.72, 1.46 0.71, 1.46 0.72, 1.51 Q4 Q4 1.74 1.22 1.15, 2.31 1.02, 2.09 1.01, 2.10	0.753
Model 1*	1.00	0.79	0.55, 1.13	0.84	0.59, 1.19	0.87	0.61, 1.24	0.676
Model 2†	1.00	0.78	0.54, 1.12	0.84	0.59, 1.19	0.87	0.61, 1.24	0.706
Model 3‡	1.00	0.85	0.59, 1.23	0.88	0.61, 1.26	0.92	0.65, 1.31	0.822
	Q1 Q2		Q2		Q3		Q4	P _{for tren}
Mean	0.01		0.07		0.13		0.30	
SD	0.04		0.01		0.02		0.14	
Crude	1.00	0.89	0.62, 1.29	1.22	0.87, 1.73	1.06	0.75, 1.52	0.499
Model 1*	1.00	0.84	0.58, 1.22	1.17	0.83, 1.65	1.03	0.72, 1.46	0.587
Model 2†	1.00	0.86	0.60, 1.24	1.17	0.82, 1.65	1.01	0.71, 1.46	0.683
Model 3‡	1.00	0.96	0.66, 1.40	1.24	0.86, 1.77	1.04	0.72, 1.51	0.704
				fgTAC	from junk food			
	Q1		Q2		Q3		Q4	P _{for tren}
Mean	0.04		0.39		0.62		1.74	
SD	0.24		0.06		0.08		1.22	
Crude	1.00	1.25	0.86, 1.81	1.01	0.68, 1.49	1.63	1.15, 2.31	0.007
Model 1*	1.00	1.15	0.79, 1.67	0.91	0.61, 1.35	1.46	1.02, 2.09	0.031
Model 2†	1.00	1.15	0.79, 1.69	0.91	0.61, 1.35	1.46	1.01, 2.10	0.034
Model 3‡	1.00	1.31	0.88, 1.95	1.06	0.69, 1.62	1.57	1.08, 2.26	0.020

fgTAC, Total Antioxidant Capacity of food groups.

* Adjusted - Sex and age.
 * Adjusted - Sex and age.
 * Adjusted - Model 1 + smoking status (never, current, former), alcohol consumption (BINGE frequency) and vitamin D consumption (mcg).
 * Adjusted - Model 2 + marital status (single/married or stable union/separated or divorced or widowed), skin colour (white and not white), physical activity (inactive/insufficiently active/active), use of supplements (yes or no), energy intake (continuous, kcal/d), baseline BMI (continuous kg/m²), professional situation.

				fgTAC f	rom natural juices			
	Q1		Q2		Q3		Q4	P _{for trend}
Mean	0.02	0.26	0.55, 1.11 0.52, 1.05 0.52, 1.05 0.57, 1.19 Q2 2.25 0.48 0.61, 1.25 0.59–1.21 0.58, 1.18 0.59, 1.22 Q2 Q2 0.05 0.01	0.54		1.33		
SD	0.13	0.05		0.12		0.75		
Crude	1.00	0.78	0.55, 1.11	0.80	0.57, 1.12	0.71	0.50, 1.01	0.099
Model 1*	1.00	0.74	0.52, 1.05	0.78	0.55, 1.10	0.70	0.49, 0.99	0.100
Model 2†	1.00	0.74	0.52, 1.05	0.78	0.55, 1.09	0.70	0.50, 1.00	0.114
Model 3‡	1.00	0.82	0.57, 1.19	0.85	0.60, 1.22	0.77	0.54, 1.11	0.235
				fgTAC fro	fgTAC from teas and coffee			
	Q1		Q2		Q3	Q4		P _{for trend}
Mean	0.42		2.25		4.19		9.14	
SD	0.68		0.48		0.72		3.90	
Crude	1.00	0.88	0.61, 1.25	0.89	0.63, 1.27	1.01	0.71, 1.42	0.807
Model 1*	1.00	0.85	0.59-1.21	0.91	0.63, 1.30	1.06	0.74, 1.51	0.533
Model 2†	1.00	0.83	0.58, 1.18	0.88	0.61, 1.27	1.02	0.71, 1.45	0.678
Model 3‡	1.00	0.85	0.59, 1.22	0.93	0.64, 1.34	1.03	0.72, 1.48	0.636
				fgTAC from artificial juices and sodas				
	Q1		Q2		Q3		Q4	P _{for trend}
Mean	0.01		0.05		0.16		0.72	
SD	0.04		0.01		0.05		0.53	
Crude	1.00	1.00	0.70, 1.45	1.00	0.70, 1.43	1.12	0.79, 1.58	0.443

Table 5. Hazard ratios and 95 % CI of depression incidence according to consumption of fgTAC by beverages, CUME Project (n 2572) (Mean values and standard deviations)

Table 5. (Continued)

				fgTAC from a	tificial juices and soc	las		
	Q1		Q2		Q3		Q4	P _{for trend}
Model 1*	1.00	0.94	0.66, 1.36	0.93	0.65, 1.34	1.10	0.78, 1.55	0.389
Model 2†	1.00	0.95	0.66, 1.38	0.94	0.65, 1.35	1.09	0.77, 1.55	0.426
Model 3‡	1.00	1.09	0.73, 1.63	1.02	0.69, 1.51	1.12	0.79, 1.61	0.568
				fgTAC from				
	Q1	Q2			Q3	Q4		P _{for trend}
Mean	0.01		0.13		0.34		1.32	
SD	0.10		0.03		0.09		1.11	
Crude	1.00	0.82	0.57, 1.17	0.81	0.58, 1.14	0.85	0.60, 1.20	0.596
Model 1*	1.00	0.77	0.54, 1.11	0.82	0.58, 1.16	0.91	0.64, 1.29	0.996
Model 2†	1.00	0.79	0.55, 1.14	0.85	0.59, 1.23	0.98	0.67, 1.43	0.694
Model 3 [±]	1.00	0.93	0.60, 1.43	1.02	0.67, 1.55	1.16	0.75, 1.77	0.323

fgTAC, Total Antioxidant Capacity of food groups.

* Adjusted - Sex and age.

+ Adjusted – Model 1 + smoking status (never, current, former), alcohol consumption (BINGE frequency) and vitamin D consumption (mcg).

‡ Adjusted – Model 2 + marital status (single/married or stable union/separated or divorced or widowedd), skin colour (white and not white), physical activity (inactive/insufficiently active/active), use of supplements (yes or no), energy intake (continuous, kcal/d), baseline BMI (continuous kg/m²), professional situation.

monitoring⁽⁵²⁾. In the same way, two cross-sectional articles, one with climacteric women and another with sixty Iranian men, found no association between dTAC, depressive symptoms or diagnosed depression^(53,54). Contrary to our findings, three cross-sectional Iranian studies observed positive associations between dTAC and the prevalence of depressive symptoms^(17–19). In a recent systematic review, our group analysed existing studies that linked dTAC and depression, concluding that consumption of an antioxidant-rich diet characterised by high dTAC scores appears to be inversely associated with depression, anxiety and sleep disorders. However, we emphasise that there are few studies available in the literature, and most have a cross-sectional design and methodological limitations, as they were conducted with Iranian individuals and, most of them, with women⁽¹⁶⁾.

Contrary to our expectations, we did not find associations between the fgTAC of the fruit and vegetables group and the incidence of depression. On the other hand, for the fgTAC of the natural juices group, the inverse association with depression remained only for the first adjustments (sex, age), not being maintained for total adjustments. A longitudinal study with Add Health Study data, which monitored 3696 17-year-old participants for 12 years, found no association between fruit and vegetables consumption and the incidence of depression either⁽⁵⁵⁾. In turn, another longitudinal study with 8353 Canadians that observed inverse associations between fruit and vegetable consumption and depression being attenuated after adjusting variables such as smoking and physical activity⁽⁵⁶⁾. Although the results of the consumption of fruits and vegetables are contradictory, a meta-analysis with observational studies found a reduction in the risk of depression with the increase in the consumption of fruits and vegetables⁽⁵⁷⁾. It is worth mentioning that fruits and vegetables and natural juices are sources of antioxidants that modulate oxidative stress, and their consumption is related to mental health^(56,57). In addition, they are related to healthier eating patterns⁽⁴⁵⁾. However, the association between the consumption of fruits and vegetables

and other behavioural factors in the incidence of depression can be complex⁽⁵⁶⁾. Thus, some behavioural factors such as physical activity, use of supplements, BMI and professional situation may have a more important impact on depression when compared with, fruits, vegetables and natural juices consumption.

The strengths of this study are its prospective design and the use of the quantitative FFQ previously validated for the study population, with good validity and reproducibility, ensuring good consistency in food consumption analyses⁽²³⁾. In addition, we highlight that the self-report of depression was previously validated for the study population⁽²²⁾. Another point is the high level of education of the participants, which can result in more reliable answers and greater adherence to the study⁽⁵⁸⁾. Finally, we highlight the use of several confounding factors for our adjustments, carefully chosen after literature reviews and with the help of a directed acyclic graph^(34,35).

As limitations, we highlight that although the FFQ has good reproducibility, we cannot guarantee that the baseline FRAP values represent the habitual long-term dietary intake precisely. Another point is the lack of national tables for FRAP values, requiring the use of values arranged in international tables for most calculations. Such factors may mediate the results observed here. Nor can we discard the possibility of residual confounding by some unmeasured or not precisely measured factors. We highlight the non-assessment of plasma TAC, but it is worth mentioning that plasma TAC may not be reflected in long-term diets, which limits its comparison with dTAC^(14,59). In addition, dTAC proves to be a handy tool in assessing the relationship between diet and health outcomes^(14,59). Finally, although the collection of data from the follow-up questionnaire Q_4 was carried out in the initial months of the COVID 19 pandemic, we cannot guarantee that this short period of time had an influence on the increase in medical diagnosis of depression. However, future analyses carried out with the next years of follow-up of the cohort may provide answers about the impact of the pandemic on the incidence of depression.

Conclusion

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Our findings do not support an association between dTAC and the incidence of depression after an average of 2.96 (1.00) years of follow-up in a highly educated Brazilian population. However, the inverse association of fgTAC from beans and lentils and the direct association of junk food with the incidence of depression in the population indicate that not only the presence of antioxidants but also the food matrix in which these antioxidants are inserted should be considered to explain the associations between diet and health outcomes. We highlight the need for further prospective studies with different nationalities to confirm these results.

Acknowledgements

The authors wish to express our gratitude to all the participants of The Cohort of Universities of Minas Gerais. The authors thank CAPES (Ministry of Education, Brazil) for granting a doctoral fellowship to G.A.P.S and A.C.G.L. J. B., H. H. M. H. and A. M. P. are research productivity fellows of CNPq (Ministry of Science and Technology, Brazil).

This study was financed in part by the Coordenação de Aperfeiçoamento de Pessoal de Nível Superior – Brasil (CAPES) – Finance Code 001. Minas Gerais Research Foundation (FAPEMIG) for the grants has supported the CUME Project (CDS-APQ-00571/13, CDS-APQ-02407/16 and CDS-APQ-00424/17).

Conceptualisation, methodology, formal analysis, investigation, data curation and writing, G. A. P. S., H. H. M. H., A. C. G. L., A. M. P., J. B., A. P. B. M. and A. S. A. All authors read and approved the final manuscript. All authors critically reviewed the manuscript and approved the final version submitted for publication.

The authors declare no conflict of interest.

Supplementary material

For supplementary materials referred to in this article, please visit https://doi.org/10.1017/S0007114523000181

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