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RESEARCH ARTICLE

Nutritional status, nutrient imbalances, food-related behaviors and dietary supplements use among patients with celiac disease in Lebanon: a national cross-sectional study

[version 1; peer review: 3 approved with reservations]

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Abstract

Background: Celiac disease (CD) is an autoimmune disorder triggered by gluten, that occurs in susceptible individuals and is associated with dietary restriction and subsequent nutritional deficiencies. This study investigated the diet quality, nutrition imbalances and nutrition status among young children, adolescents and adults with CD who were referred to several hospitals in Lebanon.

Methods: A cross-sectional study in 50 individuals (31.74 ± 15.64 years) with CD was conducted, using biochemical parameters, anthropometric measurements, dietary and physical activity assessments.

Results: Of the 50 participants, 38% and 16% were presenting low serum levels of iron and vitamin B12, respectively. The majority of participants were physically inactive and around 40% of them had low muscle mass. A weight loss of 10% to 30% indicating mild to moderate

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malnutrition was shown in 14% of individuals. The assessment of food-related behaviors shows that 80% of participants were reading nutrition labels and 96% of them were following gluten-free diets (GFD). Some barriers including family ignorance (6%), language of the nutrition labels (20%) and expensive GF products (78%) were limiting the adherence to GFD. The inadequacy of the daily energy intake along with insufficient intakes of calcium and vitamin D were remarked among individuals with CD. However, protein and iron intake were exceeding the recommendations among all age groups, except in males aged 4-8 years and 19-30 years. Half the study participants were using dietary supplements where 38%, 10%, 46%, 18%, 16% and 4% used vitamin D, vitamin B12, iron, calcium, folate and probiotics, respectively.

Conclusion: GFD is the key treatment for CD. However, it is not without inadequacies and may cause certain deficiencies such as calcium and vitamin D leading to reduced bone density. This underlines the critical role of dietitians in education and maintenance of healthy GFD among individuals with CD.

Keywords


Celiac disease, gluten free diet, anthropometry, biochemical, Lebanon





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Introduction

Celiac disease (CD) is a chronic inflammatory autoimmune disorder characterized by villous atrophy of the small bowel.¹ The exposure to dietary wheat gluten and proteins of rye and barley triggers CD among genetically predisposed persons.² In certain circumstances, individuals show no symptoms and the diagnosis is only possible through serologic screening.³ According to the European Society for the Study of Celiac Disease (ESsCD) guideline, the diagnostic criteria depend upon the exposure to dietary gluten or related proteins of rye and barley-dependent symptoms, through some histopathologic findings (villous atrophy and crypt hyperplasia) from a biopsy of the duodenum CD, specific antibody levels, HLA-DQ2 and/or HLA-DQ8.⁴ The prevalence of CD is estimated to be one to two individuals per 100 worldwide⁵ and roughly one in 99 among children and adolescents.⁶ Moreover, according to a recent systematic review and meta-analysis, the pooled seroprevalence of CD in the general population is significantly higher in children (2%) compared with adults (1%, p-value=0.01) and significantly greater among females (1.65%) compared to males (0.8%, p-value=0.04).⁷ The pooled sero-prevalence of CD in the Middle Eastern area (1.47%) is higher than that in South Asia (1.25%) and East-Asia (0.06%).⁷ The highest prevalence among the Arab regions was reported in Saudi Arabia (3.2%), and the lowest in Tunisia (0.1%).⁸ In Lebanon, the prevalence of CD was around 0.5% in 2011.⁹ Furthermore, women have shown elevated prevalence compared to men in the Arab countries⁸ and according to solid evidence, the risk becomes common and high if a first- or second-degree relative has been previously diagnosed.⁶

CD affects the proximal part of the small intestine, which in turn results in micronutrient malabsorption, particularly iron, folic acid, vitamin B12, calcium and vitamin D, resulting in anemia on the short term and reduced bone density on the long-term.² Further, beyond inflammation and malabsorption, CD is characterized by recurrent abdominal pain, nausea, vomiting, steatorrhea and loss of weight which also compromise the nutrition status of affected individuals.⁶ Until today the only therapy for individuals with CD remains the adherence to a gluten-free diet (GFD) along with the avoidance of foods containing wheat, rye, and barley derivatives.¹⁰ According to several studies, the adherence to a strict GFD can reduce the risk of health-related complications, for instance: malignancy, osteoporosis, growth retardation in children, liver disorders and reproductive tract disorders.¹ According to recent data, the adherence to GFD has remained unchanged over the past twenty years^{11,12} and a considerable number of individuals with CD do not adhere to the dietary restrictions, which affects the patient's quality of life and causes compromised nutrition status.¹⁰ Moreover, some individuals with CD may look for other approaches to treatments such as complementary and alternative medicine (CAM) which include dietary supplements (DS) use.¹³ There is scarce data on the DS use in CD. Many studies identified the DS use among CD patients to be within a range of 21.6% to 23.6%.¹³ This percentage was less than the percentage of DS use among the general population (27%).¹³ Thus, investigating the nutrition status through dietary assessment of individuals with CD is warranted especially among children, adolescents, and adults to avoid malnutrition and related morbidity. Such an evaluation of GFD adherence should be executed by experienced trained dietitians; however, the access to such dietitians with expertise in the field of CD may be limited. Beyond GFD adherence, many reasons exist for the necessity of nutritional investigation including the poor public awareness about CD, and the lack of regulations for the production and analysis of gluten-free labelled products.¹⁴ To the best of our knowledge, there are no published data that address the nutrition situation and dietary supplements use among individuals with CD in Lebanon. Therefore, the aim of the present study was to assess the nutritional status, food-related behaviors, caloric, macro- and micronutrients intake and dietary supplements use among individuals with CD in Lebanon.

Methods

Study design and participants

Selection procedure

Participants from all age categories were invited to participate in the current cross-sectional study between February and July 2018. A random selection was performed. Participation in the study was proposed to all patients who came to the pediatrician-gastroenterologists and gastroenterologists in three hospitals in Beirut and Mount Lebanon for a follow-up visit and met the criteria for inclusion, which was a diagnosis of CD based on serological and histological markers according to the international guidelines,¹⁵ more than six months prior to study enrolment to guarantee adequate GFD knowledge. All participants with no proven biopsy, pregnant women, history of cancer, eating disorders, malnutrition, renal diseases and any other factors that affect the nutrition situation, were excluded.

Questionnaires and dietitians' assessment

The survey contained 78 questions concerning demographics, disease's characteristics, history and presentation, food-related behaviors, lifestyle and diet history, weight before and after being diagnosed with CD, adherence to GFD and social and emotional barriers for sustaining the GFD. The questionnaire was derived from the one available from the

Canadian Celiac Association [website](#) and used previously in the “Canadian Celiac Health Survey”.¹⁶ Medical history was evaluated based on the age at diagnosis of CD, family history of CD, symptoms (diarrhea, headache, flatulence and bloating), comorbid diseases (food allergies, food intolerance, other non-communicable disease). Alongside the first questionnaire, additional FFQ for caloric and protein intake, calcium, vitamin D and iron intake was addressed to individuals with CD along with questions on dietary supplement use. This FFQ was derived from two Lebanese-validated questionnaires to address the questions for children, adolescents and adults^{17,18} in which a substitution of 24 foods with GF foods was executed (see *Underlying data*⁶⁹). Dietary supplement use was assessed through patient’s response of “Yes” or “No” to the statement: “I use dietary supplements in addition to GFD”. To investigate the type of supplements used, patients were asked to answer the following question: “Since you were diagnosed, have you used any of the following dietary supplements?” The specific options included vitamin D, probiotics, iron, folate, calcium, and vitamin B12, which are the commonly used DS among individuals with CD.¹⁹ The questionnaire used in the present study was reviewed by two national and one international expert in CD and was pre-tested in 14 non-expert individuals as a focus group. The aim of the focus group was to explore a specific set of issues: the questionnaire’s readability, reliability and inter consistency (Cronbach-alpha=0.69). Moderators often commence the focus group by asking broad questions about the topic of interest, before asking the focal questions. Although participants individually answer the facilitator’s questions, they are encouraged to talk and interact with each other. In addition, three-day food records were collected and analyzed using *Nutritionist Pro* software (version 3.2, AXXYA) to assess caloric, micro- and macronutrients intake. The mean value and the standard deviation (SD) were used when interpreting the nutrient’s intake.

Weight and height were measured using automated scale and stadiometer. Body composition including body fat, lean body mass and visceral fat were assessed using Bio-impedancemetry body composition analyzer (BF-511, OMRON). The standard formula of body mass index (BMI) was calculated by dividing weight in kilograms (kg) by height in meters squared (m²).

Laboratory values

According to a study by Bledsoe *et al.* (2019),¹⁹ and Marteau *et al.* (2001)²⁰ albumin and micronutrient deficiencies, including vitamins B12 and D, as well as folate, iron, zinc and copper, are common in adults at the time of diagnosis with CD. Thus, the study participants were tested for three out of six micronutrients (serum iron, folate and vitamin B12) along with albumin levels in the present study. Venous blood was collected after a 12-hour period of fasting, centrifuged and stored at –80°C. The iron level was measured in serum by the *in vitro* assay method with Roche Diagnostic (Mannheim, Germany) and Hitachi 704 devices (Roche Diagnostics, Switzerland).

All patients underwent a complete hemogram. However, only albumin, serum iron, folate and vitamin B12 were reported in this study to comply with the outcomes discussed in Bledsoe *et al.*¹⁹ Serum iron was diagnosed according to the criteria followed by the laboratory in which the analysis occurred.²¹ The reference values for children were: 8.9–21.4 µmol/L (SI units), males (adults): 14–32 µmol/L (SI units) and females (adults): 11–29 µmol/L (SI units) were compared to the values obtained in the current study. Serum folate and vitamin B12 levels were measured using a competitive immunoassay. As for vitamin B12, the reference values for 1–<11 years: 260–1200 pmol/L, 12–<18 years: 200–800 pmol/L, 18–<65 years: 200–600 pmol/L²² were also used to compare the results obtained in this study. Concerning the folate reference values, the normal serum levels were, for children: 5–21 ng/ml and for adults: 2–20 ng/mL.²³ As for albumin, it was considered low if the value was <3.5 and high if >5.5 g/dL.²⁴

Food-related behaviors and GFD dietary adherence

Some food patterns including shopping, cooking and eating out habits were assessed among the study participants. The adherence to GFD was assessed by trained dietitians who investigated the infringement regarding dietary recommendations.¹² Across the three-day food records, patients presenting a consumption between 3 to 10 g of gluten per day were considered as non-adherent.^{12,25}

Facilitators and barriers

To search for possible barriers, patients were asked, “what are the major challenges that limit the adherence to a strict gluten-free diet?”. Trained dietitians asked participants to report whatever came to mind concerning this question. The answers were analyzed according to the methodology used by Braun and Clarke.²⁶

Indicators of malnutrition and criteria for the determination of nutritional status

The determination of the nutritional status was based on anthropometric data (weight, height, BMI), hypoalbuminemia and the percentage of weight loss taking into consideration age and sex factors. A weight loss of 10% to 20% indicated mild malnutrition, a weight loss of 20% to 30% characterized moderate malnutrition, and a weight loss of more than 30% indicates severe malnutrition.²⁵

Physical activity

Several types of physical activity such as aerobic (fitness), swimming, running, strength training gym, walking, cycling and yoga were investigated. The frequency of physical activity was based on frequency per week: one time per week, two-three times per week, four or more per week, daily or no physical activity.

Statistical analysis

Statistical analysis of the study was performed using SPSS 28.0 (IBM Statistical Package for Social Sciences software version 28). The descriptive data were presented as percentages for categorical variables and mean \pm SD for quantitative variables. Categorical variables were compared with the Chi-square test and continuous variables were compared with the student-T test. $P < 0.05$ was considered a statistically significant difference with a 95% confidence interval.

Ethical considerations

At the beginning of the study, a consent form was signed by participants. Moreover, this study received the approval of the ethics committees at the Lebanese university (CU#18-46). In addition, an assent form was signed by the parents of children and adolescents under 18 years old.

Results

A total of 50 participants (78% females; mean age = 31.74 ± 15.64 years) was enrolled in the present study. Out of 50, eight participants were aged below 18 years. Age categories were classified according to the dietary reference intake classes²⁷ of which 46% of the population's age ranged between 31 and 50 years. In the current study, females were younger than males (p -value = 0.03). Moreover, around 66% of all the participants had university degrees and more than 70% were remunerated for more than 1,500 million Lebanese pounds (LBP) per months. Half the study's participants were married. Overall, in almost all the characteristics, there were no significant differences between females and males in this study ($p > 0.05$). The sociodemographic characteristics of the study participants are displayed in [Table 1](#).

Table 1. Socio-demographic characteristics of biopsy-proven study population, overall and by gender.

LBP: Lebanese Pounds.

Variables	Overall (N=50) mean (SD)/N (%)	F (N=39;78%) mean (SD)/N (%)	M (N=11; 22%) mean (SD)/N (%)	p-value
Age (years) Mean \pm SD ^a	31.74 \pm 15.64	34.15 \pm 2.50	23.18 \pm 3.58	0.03
Age groups ^b				
4-8	5.5 \pm 1.64/6(12)	6.0 \pm 0.91/4(10.5)	4.5 \pm 0.5/2(18.2)	0.37
9-13	12.6 \pm 1.15/2(4)	13 \pm 1.0/1(2.6)	12 \pm 0.0/1(9.1)	0.66
14-18	-	-	-	-
19-30	24.5 \pm 3.34/12(24)	25 \pm 1.17/7(18.4)	23 \pm 1.52/5(45.5)	0.36
31-50	38.08 \pm 5.6/23(46)	38.3 \pm 1.59/20(52.6)	37.6 \pm 1.67/3(27.3)	0.78
51-70	57.5 \pm 5.2/7(14)	58.25 \pm 3.19/6(15.8)	56 \pm 2.0/0(0)	0.67
Income level per month (%)				0.47
\geq 1,500,000 LBP	36 (72)	29 (74.4)	7 (63.6)	
<1,500,000 LBP	14 (28)	10 (25.6)	4 (36.4)	
Education level (%)				0.64
Illiterate	0 (0)	1 (2.6)	0 (0)	
School	11 (22)	9 (23.1)	2 (18.2)	
High school	6 (12)	5 (12.8)	1 (9.1)	
University	33 (66)	24 (61.5)	8 (72.7)	

Table 1. *Continued*

Variables	Overall (N=50) mean (SD)/N (%)	F (N=39;78%) mean (SD)/N (%)	M (N=11; 22%) mean (SD)/N (%)	p-value
Marital status (%)				0.6
Single	23 (46)	17 (43.6)	6 (54.5)	
Married	26 (52)	21 (53.8)	5 (45.5)	
Divorced	1 (2)	1 (2.6)	0	

^aSD=standard deviation.^bclassification according to the DRIs values.²⁷

Table 2 includes the medical history, anthropometric measures and lifestyle characteristics of the study participants. In the current study, males were taller than females (p-value=0.042). It was observed that the weight of all participants increased in a range of 4 to 5.6 kg since being diagnosed with CD. This was significantly higher among males compared to females

Table 2. Anthropometry measures, medical and lifestyle characteristics of the study population, overall and by gender. CD: celiac disease; BMI: body mass index.

Variables	Overall (N=50) mean (SD)/N (%)	F (N=39;78%) mean (SD)/N (%)	M (N=11; 22%) mean (SD)/N (%)	p-value ^d
Anthropometry indices				
Weight (kg)				
Weight before GFD ^a	59.99±2.69	41.54±6.22	57.92±4.97	0.025
Current weight	64.31±3.27	44.02±7.06	63.60±5.06	0.017
Weight change ^b	4.32±1.84	3.93±2.29	5.68±2.03	0.023
Risk of malnutrition (weight loss in the past 6 months)				0.31
No risk (<10%)	43 (86)	32 (82.1)	11 (100)	
Mild (10%-20%)	6 (12)	6 (15.4)	0 (0)	
Moderate (20%-30%)	1 (2)	1 (2.6)	0 (0)	
High risk (>30%)	0 (0)	0 (0)	0 (0)	
Height (cm)	159.56±18.62	145.45± 7.04	161.9±8.23	0.042
BMI adults ^c (≥19 years)				
<18 kg/m ²	2 (5)	2 (6.1)	0 (0)	0.66
18-25 kg/m ²	21 (52)	16 (48.5)	5 (62.5)	
≥25 kg/m ²	18 (43)	15 (45.5)	3 (37.5)	
Muscle mass (kg) (≥6 years) ²⁸	27.31±7.03	26.21±0.72	31.47±3.9	0.034
Low	20 (40)	15 (39.5)	5 (50)	
Normal	23 (46)	23 (60.5)	0 (0)	
High	1 (2)	0 (0)	1 (10)	
Very high	4 (8)	0 (0)	4 (40)	
Fat mass (kg) (≥6 years) ²⁸	32.21±16.6	35.18±2.75	20.94±2.80	0.014
Low	2 (4)	1 (2.6)	1 (10)	
Normal	16 (32)	13 (34.2)	3 (30)	
High	13 (26)	10 (26.3)	3 (30)	
Very high	17 (34)	14 (36.8)	3 (30)	
Visceral mass (kg) (≥6 years) ²⁸	6.59±5.9	6.06±0.4	8.61±4.15	0.254
Normal	40 (80)	32 (94.1)	8 (100)	
High	2 (4)	2 (5.9)	0 (0)	

Table 2. Continued

Variables	Overall (N=50) mean (SD)/N (%)	F (N=39;78%) mean (SD)/N (%)	M (N=11; 22%) mean (SD)/N (%)	p-value ^d
Medical history				
Family history of CD				
No	29 (58)	23 (59)	6 (54.5)	0.79
Yes	21 (42)	16 (41)	5 (45.5)	
Age at diagnosis	22.28±13.55	24.34±13.84	15.18±10.01	0.26
Symptoms				
No	3 (6)	1 (2.6)	2 (18.2)	0.054
Yes	47 (94)	38 (97.4)	9 (81.8)	
Severity of symptoms				
Mild	17 (34)	12 (30.8)	5 (45.5)	0.32
Moderate	16 (32)	12 (30.8)	4 (36.4)	
Severe	17 (34)	15 (38.5)	2 (18.2)	
Related illness				
No	38 (76)	29 (74.4)	9 (81.8)	0.609
Yes	12 (24)	10 (25.6)	2 (18.2)	
Medications				
No	39 (78)	29 (74.4)	10 (90.9)	0.242
Yes	11 (22)	10 (25.6)	1 (9.1)	
Laboratory values				
Albumin (3.5-5.5 g/dl) ²⁴	4.41±1.05	4.57±0.72	3.84±1.72	0.02
Low	7 (14)	2 (5.1)	5 (45.5)	
Normal	36 (72)	32 (82.1)	4 (36.4)	
High	7 (14)	5 (12.8)	2 (18.2)	
Iron (μmol/L (SI units) ²¹	31.83±10.49	30.80±9.84	35.50±12.35	0.40
Low	19 (38)	15 (38.5)	4 (36.4)	
Normal	20 (40)	14 (35.9)	6 (54.5)	
High	11 (22)	10 (25.6)	1 (9.1)	
vit. B12 (pmol/l) ²²	329.52±110.8	323.26±18.7	351.72±26.29	0.10
Low	8 (16)	8 (20.5)	0 (0)	
Normal	42 (84)	31 (79.5)	11 (100)	
High	0 (0)	0 (0)	0 (0)	
Folate (ng/mL) ²³	14.79±3.36	14.51±3.35	15.78±3.38	0.273
Low	0 (0)	0 (0)	0 (0)	
Normal	45 (90)	35 (89.7)	10 (90.9)	
High	3 (6)	2 (5.1)	1 (9.1)	
Lifestyle				
Smoking				0.78
No	39 (78)	31 (79.5)	8 (72.7)	
Yes	11 (22)	8 (20.5)	3 (27.3)	
Alcohol (per week)				
No	41 (82)	30 (78.9)	10 (90.9)	0.36
Yes	9 (18)	8 (21.1)	1 (9.1)	

Table 2. Continued

Variables	Overall (N=50) mean (SD)/N (%)	F (N=39;78%) mean (SD)/N (%)	M (N=11; 22%) mean (SD)/N (%)	p-value ^d
Physical activity (per week)				
No	36 (72)	29 (74.4)	7 (63.6)	0.69
Yes	14 (28)	10 (25.6)	4 (36.4)	
<1 time per week	4 (28.6)	4 (40)	0 (0)	0.13
2-3 times per week	10 (71.4)	6 (60)	4 (100)	
>4 times per week	0 (0)	0 (0)	0 (0)	

^aGluten free diet.^bWeight change is calculated by subtracting weight after from the weight before and the significance is determined by paired sample t-test.^cSubjects aged under 20 years were excluded from this analysis.^dp<0.05 indicates the significance of values of independent tests between the means of female and males.

(p-value=0.023). Moreover, 45% of females were overweight compared to 37% males. The assessment of malnutrition was based on Kotze *et al.*²⁵; a weight loss of 10% to 30% indicating mild to moderate malnutrition were shown in 14% of the study population. The body composition analysis conducted for the study participants revealed that 40% of them had low muscle mass (5.0 to 32.8% for males and 5.0 to 25.8% for females).²⁸ Further, the mean muscle mass was higher in males compared to females (p-value=0.034). In total, 60% of participants were having high fat mass (more than 20% in males and more than 30% in females). In addition, the mean value of fat mass among women were higher than that among men (p-value=0.014). Furthermore, the majority of the study participants had low visceral fat and there were no significant differences regarding the mean and SD of visceral fat among both genders (p-value=0.25).

As for the medical history, around half the study participants had a family history of CD. The mean age at diagnosis was 22.28±13.55 (range=1-57 years). Symptoms related to CD such as abdominal bloating, diarrhea, headache, and flatulence appeared in most participants (94%) and 6% were asymptomatic. Around 66% were facing moderate to severe symptoms. The participants were asked about comorbidities along with CD; only 24% of them had CD with other diseases of which dermatitis, arthritis, lactose intolerance and food allergy. As for the medications, 78% of the study participants reported being on medical treatments for CD symptoms.

As for the laboratory values, hypoalbuminemia was observed in 14% of the study population. Women were having significantly higher mean and SD levels of albumin compared to men (p-value=0.02). Out of 50 participants, 38% and 16% were presenting low serum levels of iron and vitamin B12, respectively. None of the study participants presented with folate deficiency. As for the lifestyle behaviors, the majority of participants were non-smokers, non-drinkers and physically inactive (Table 2).

The food-related behaviors assessed among the study participants are presented in Table 3. Around 80% of the study participants reported reading nutrition labels when shopping groceries. Moreover, almost 96% of them were following a nutritional diet of which was the GFD. To assess their adherence to GFD, the examination of 96% of the participant's three-day records reported low intake of gluten (<3g per day). Only 4% weren't adhering to the GFD recommendations. As reported by the participants, some barriers including family ignorance (among 6% of the population), language on the nutrition labels (among 20% of the population) and expensive GF products (among 78% of the population) were limiting the adherence to GFD. It was observed that 72% of the study population had good cooking experience and 64% of the study participants were involved in preparing meals for themselves. On the other hand, almost all the subjects were unaware of a possible cross-contamination during cooking. Despite that, 82% of the subjects with CD preferred fast foods, although they consumed their meals frequently at home (92%). It was reported that the adherence to GFD by these participants induced a hunger feeling during the day. Most of our study population necessitated extra care by family members regarding cooking and eating. No significant differences were observed between genders.

Energy and protein intake compared to the recommended daily allowances (RDA)

Table 4 shows the mean energy and nutrients intake calculated through the three-day food records reported by the study participants or their families. Among female children aged four to eight years, the mean daily energy (E) and protein (P) intake were estimated as 1396 ±424.15 Kcal/d and 56.41 ±18.31 g/d, respectively. In comparison with the reference values of the recommended daily allowances (RDA), the daily intake of E and P was 130% and 268%, respectively. On the other hand, it was observed that female children aged 9-13 years had insufficient daily E intake (75% vs RDA) but

Table 3. Food-related behaviors among the study population, overall and by gender. GF: gluten-free; GFD: gluten-free diet; F: female; M: male.

Outcome	Overall (N=50) N (%)	F (N=39;78%) N (%)	M (N=11; 22%) N (%)	p-value
Reading labels when eating and shopping				
No	10 (20)	6 (15.4)	4 (36.4)	0.12
Yes	40 (80)	33 (84.6)	7 (63.6)	
Following diet				
None	2 (4)	2 (5.2)	1 (5.22)	0.32
GF diet	48 (96)	37 (94.9)	10 (90.9)	
Adherence to GFD^a				0.06
No	1 (2)	0 (0)	1 (9.1)	
Yes	48 (96)	38 (100)	10 (90.9)	
Family support in GF diet's adherence				0.63
No	3 (6)	3 (7.7)	0 (0)	
Yes	47 (94)	36 (92.3)	11 (100)	
Language of food label as barriers in GF diet's adherence				0.70
No	40 (80)	32 (82.1)	8 (72.7)	
Yes	10 (20)	7 (17.9)	3 (27.3)	
Financial barriers towards GF diet's adherence				0.84
No	11 (22)	9 (23.1)	2 (18.2)	
Yes	39 (78)	30 (76.9)	9 (81.8)	
Cooking experience				0.026
No	14 (28)	8 (20.5)	6 (54.5)	
Yes	36 (72)	31 (79.5)	5 (45.5)	
Self-preparation of meals				
No	18 (36)	12 (30.8)	6 (54.5)	0.147
Yes	32 (64)	27 (69.2)	5 (45.5)	
Awareness of cross-contamination during meal preparation				
No	49 (98)	38 (97.4)	11 (100)	0.592
Yes	1 (2)	1 (2.6)	0 (0)	
Fast foods vs home meals				
Fast foods	41 (82)	32 (82.1)	9 (81.8)	
Home meals	9 (18)	7 (17.9)	2 (18.2)	0.986
Eating out vs. eating at home				
Home	46 (92)	37 (94.9)	9 (81.8)	0.159
Eating out	4 (8)	2 (5.1)	2 (18.2)	
Feeling hungry most of the time				0.08
No	14 (30)	14 (35.9)	1 (9.1)	
Yes	25 (70)	25 (64.1)	10 (90.9)	
Looking for extra care in meals preparation				
No	10 (20)	8 (20.5)	2 (18.2)	0.847
Yes	40 (80)	31 (79.5)	9 (81.8)	

^aPatients presenting an ingestion of approximately 3–10 g of gluten per day were considered as non-adherent.^{12,25}

Table 4. Energy and protein intake and their reference values according to recommended daily allowance (RDA) among the study population. EI: energy intake; PI: protein intake.

Variables	Overall	Female	RDA	% of adequacy	Male	RDA	% of adequacy	p-value (F vs M)
4-8								
EI (kCal)	1396±424.15	1564±88.6	1200	130	1060±483	1400	75	0.195
PI (g)	56.41±18.31	51.22±9.8	19	268	14.14±10	19	74	0.384
9-13								
EI (kCal)	1202.5±272.2	1010±0.0	1600	75	1395±0.0	1800	77.5	0.024
PI (g)	39.85±39.85	108.36	34	317	52.00	34	152	0.047
19-30								
EI (kCal)	1564±177.26	2076.25±39.7	2200	94	1530±255.2	2900	52.7	0.024
PI (g)	76.84±35.4	84.97±8.86	46	184	92.7±12.92	56	60	0.047
31-50								
EI (kCal)	1758.3±772.4	1669.85±177.1	2200	76	2348±26.1	2400	98	0.942
PI (g)	178±35.43	71.99±9.702	46	156	84.73±11.5	56	151.8	0.423
50-70								
EI (kCal)	1701.08±296	1701.08±296	1900	89.5	0.024	2300	-	0.024
PI (g)	81.183±33	81.183±33	46	176	0.047	56	-	0.047

*The analysis couldn't be generated due to the low sample size; the missing values in the table are due to the absence of males above the age of 50 years.

Table 5. Average daily nutrient intake of subjects and percentage of adequacy according to recommended daily allowance (RDA).

Nutrient intake/Age	Mean of overall population	Female			Male			P-value (F vs M)
		Actual intake	RDA ²⁹	% Adequacy	Actual intake	RDA ²⁹	% Adequacy	
Calcium (mg/day) ^{a,b}								
4-8	702.41±118.07	768.63±173.75	1000	77%	569.98±41.02	1000	57%	0.98
19-30	540.16±325.47	601.92±404.41	1000	60%	453.96±173.13	1000	45%	0.46
31-50	536.34±287.18	547.36±306.84	1000	55%	462.91±58.55	1000	46%	0.64
51-70	454.78±190.89	454.78±190.89	1200	38%	-	-	-	-
Iron (mg/day) ^{a,b}								
4-8	16.303±3.11	22.48±7.48	10	194%	15.17±4.72	10	101%	0.09
19-30	30.80±5.55	33.49±6.39	18	158%	27.03±10.56	8	338%	0.59
31-50	17.22±3.95	17.58±4.49	18	98%	14.79±6.37	8	185%	0.81
51-70	18.81±7.56	18.81±7.56	8	235%	-	8	-	-
Vitamin D (ug/day) ^{a,b}								
4-8	6.54±2.21	8.56±2.87	15	57%	2.52±0.06	15	17%	0.003
19-30	10.42±4.93	8.48±2.91	15	57%	13.12±11.82	15	88%	0.66
31-50	3.56±0.89	3.94±1.00	15	26%	1.00±0.39	15	7%	0.28
51-70	2.28±0.62	2.28±0.62	15	15%	-	-	-	-

^aIn each age category, the age groups (13-14) were removed due to the limited number of patients.^bThe missing values in the table are due to the absence of males above the age of 50 years.

Table 6. Supplements intake among study participants, by age group, by gender and reference to recommended daily allowance RDA for each element.

	Overall (N=50) N (%)	F (N=39;78%) N (%)	M (N=11; 22%) N (%)	P-value
Dietary supplements use				0.108
No	23 (46)	16 (41)	7 (63.6)	
Yes	27 (54)	23 (59)	4 (36.4)	
Vitamin D				0.125
No	31 (62)	22 (56.4)	9 (81.8)	
Yes	19 (38)	17 (43.6)	2 (18.2)	
Vitamin B12				0.909
No	45 (90)	35 (89.7)	10 (90.9)	
Yes	5 (10)	4 (10.3)	1 (9.09)	
Iron				0.158
No	27 (54)	19 (48.7)	8 (72.2)	
Yes	23 (46)	20 (51.3)	3 (27.8)	
Calcium				0.384
No	41 (82)	31 (79.4)	10 (90.9)	
Yes	9 (18)	8 (20.6)	1 (9.09)	
Folate				0.261
No	42 (84)	31 (81.5)	11 (100)	
Yes	7 (16)	7 (18.5)	0	
Probiotics				0.443
No	48 (96)	37 (94.8)	11 (22)	
Yes	2 (4)	2 (5.2)	0	

surpassed by 317% the daily protein requirements. Similarly, among the female participant's age groups 19-30, 31-50 and 50-70, there were inadequacies in the daily E intake along with overconsumption of protein per day. Otherwise, inadequacies in the daily E and P intake were observed among the male children (four to eight years) and those aged 19 to 30 years. Moreover, among those aged 9-13 and 31-50, the daily E intake was inadequate, but the P intake was high. No significant differences regarding the daily E and P intakes were observed between females and males, except in the 19-30 age group, where the daily energetic and protein ratios were high in females compared to males (p-value=0.024 and p=0.047, respectively).

The average daily nutrient intake of calcium, iron and vitamin D is reported in Table 5. Among females and males, none of the study participants showed adherence to the daily recommendations for calcium and vitamin D intake. On the other hand, almost all the study participants from both genders showed a high daily consumption of iron.

Dietary supplements were used by 54% of the study population of which 38%, 10%, 46%, 18%, 16% and 4% used vitamin D, vitamin B12, iron, calcium, folate and probiotics, respectively. No difference was observed between genders regarding DS use (Table 6). According to Figure 1, it appears that the usage of DS among individuals with CD was not excessive.

Discussion

This study, the first of its kind in Lebanon, highlights the nutrition imbalances, P-E malnutrition, adherence to GFD, food-related behaviors and dietary supplements use among participants with CD. Our findings show that 78% of individuals with CD were women. This finding was higher than the data reported by Jansson-Knodell *et al.*,³⁰ and Ashtari *et al.*⁷ which showed that the prevalence of women diagnosed with CD was 65% between 1990 and 2015 and the sero-prevalence of CD reported among women was 165 women out of 2000, respectively. On the other hand, our finding aligned the data reported by Singh *et al.* which showed that the prevalence of CD was higher among women compared to men.³¹ In addition, according to Bai *et al.*, there are physiological differentials among genders in CD³²: “men have

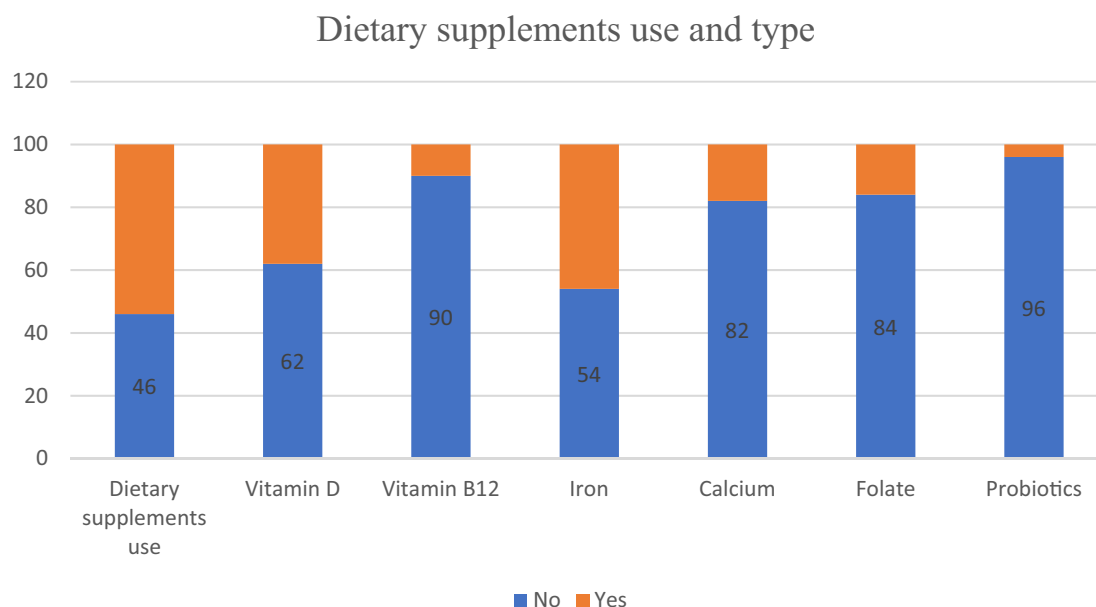


Figure 1. Dietary supplements use and type among study participants.

indirect evidence of greater malabsorption than females and have female-predominant associated diseases when they present with celiac disease³². This study supported our finding concerning the remarked P-E deficiency among males compared to females in our study. Further, almost all the participants had symptoms related to CD such as abdominal bloating, diarrhea, headache, and flatulence; around 66% were facing moderate to severe symptoms and 78% of them reported being on a medical treatment for CD symptoms. These findings were two times higher than the prevalence reported by Stasi *et al.*, who showed that 22% of individuals with CD faced persistent abdominal symptoms.³³ In the present study, 14% presented hypoalbuminemia and mild to moderate malnutrition (10%-30% weight loss in the previous six months). This data agreed with Wierdsma *et al.*³⁴ where 17% of individuals with CD were malnourished (>10% undesired weight loss). On the other hand, the percentage of underweight women (5%) in our study wasn't in accordance with Wierdsma *et al.*³⁴ in which 22% of the women were underweight and 29% of the patients were overweight (BMI > 25 kg/m²). Our study showed that around 40% of individuals with CD had low muscle mass and 60% had high fat mass. This result was in contradiction with the results reported in a national study which showed that subjects with CD had a lower fat mass (17.8±2.0 kg)³⁵ but in concordance with two recent systematic reviews on children and adults that showed that after one year follow-up to GFD' adherence, fat mass of CD patients significantly increased compared to that at baseline.^{36,37} According to Agarwal *et al.*, patients with CD are more susceptible to develop metabolic syndromes and fatty liver diseases.³⁸ Thus, these patients should be regularly counseled by experienced dietitians for nutritional and metabolic factors about healthy diets and physical activity. However, the majority of participants in our study were physically inactive (72%). This prevalence of sedentary patterns among our study population was higher than that reported by Nestares *et al.* (37%) among children and adolescents³⁹ but lower than the prevalence of physical activity among women with CD in Poland (70%).⁴⁰

Out of 50 participants, 38% and 16% were presenting low serum levels of iron and vitamin B12, respectively. None of the study participants presented folate deficiency. These results showed consistency with a recent systematic review that showed high occurrence of hypoalbuminemia and anemia among individuals with CD.⁴¹ Furthermore, micronutrient deficiencies are common features among individuals with CD. This was demonstrated in the study of Bledsoe *et al.*¹⁹ where weight loss was seen in 25.2%, albumin was low in 19.7%, vitamin B₁₂ was low in 5.3% and folate was low in 3.6% of cases. Moreover, it was shown that 87% of patients with CD in the study of Wierdsma, *et al.*³⁴ had low vitamin B₆ (14.5%), folic acid (20%), and vitamin B₁₂ (19%). Additionally, 46% of patients with CD had low serum iron, and 32% had anemia.³⁴

In our study, around 80% of participants read nutrition labels when shopping groceries. Reading and understanding food label's topics have been discussed widely and have been associated with strict GFD adherence.⁴² Our finding was in accordance with Butterworth *et al.*, who showed that amongst 74% of individuals with CD, some factors were in compliance with adherence to GFD: being a member of a celiac society, understanding food label's language, and having

access to gluten-free products with reasonable prices, regular dietetic and medical follow-up.⁴³ Causes of adherence/non-adherence to a GFD are multifactorial and differed according to their nature and magnitude.^{44–47}

Almost 96% of our population were following a GFD. To assess their adherence to GFD, the examination of the participant's three-day records reported low intake of gluten (<3g per day) among 96% of the study participants. Only 4% weren't adhering to the GFD recommendations. This finding was consistent with a study conducted in individuals with CD in Spain (92.5%).⁴⁸ According to a Spanish study, 72% of individuals with CD showed acceptable adherence which was associated with higher levels of self-efficacy.⁴⁹ Three studies from the UK showed that white patients were adhering to GFD in a range from 53 to 81%.⁵⁰ Non-adherence to GFD was also associated with diagnosis at younger ages and smoking.⁵⁰ This was not shown in our study. In Canada, GFD adherence was assessed and revealed that the percentage of strict adherence was only 56%.⁵¹ The assessment of GFD adherence of 70 Swedish adolescents with CD showed that 86% of them were adherent to GFD five years after screening.⁵² Moreover, of 5310 adult and adolescent Australians and New Zealanders with CD, 61% adhered to a GFD.⁵³ Three studies on adolescents from Brazil identified adherence rates from 36 to 86% taking into consideration that younger children (up to 12 years) were more likely to comply with the diet⁵⁴ while teenagers interrupted their diet purposefully. Parents of pediatric patients with CD from the Slovak Republic, reported a GFD adherence of 69% by children.⁵⁵ In addition, among 38 studies, the adherence was ranging from 42 and 91%.⁴⁹ According to a recent systematic review of 49 studies, the adherence rates were ranging from 23 to 98%.⁵⁶ This wide variability in adherence rates may be explained by the different populations examined (*e.g.*, adults, adolescents, children, ethnic minorities), but also by the different tools used for evaluating adherence.

As reported by the participants, some barriers including family ignorance (among 6% of the population), language on the nutrition labels (among 20% of the population) and expensive GF products (among 78% of the population) were limiting the adherence to GFD. These findings were in concordance with Muhammad *et al.*,⁴⁴ who showed that not understanding food labelling was associated with lower GF dietary adherence scores. A higher proportion of individuals in the United Kingdom (UK) reported difficulties in acquiring knowledge about GF products (ranging between 5% and 76%) and understanding of food labels (4%-53%).⁵⁷ It was observed that 72% of the study population had good cooking experience and 64% of the study participants prepared their meals. On the other hand, almost all the subjects were unaware of a possible cross-contamination during cooking food. Despite 82% of the subjects with CD preferring fast foods, they frequently consumed their meals at home (92%). Beyond cooking meals at home and cross-contamination, our findings were consistent with the findings reported in a recent systematic review indicating that the most significant barriers limiting the adherence to GFD included: "lower knowledge of CD" (35%); "restaurant/supermarket shopping" (30%); "poor patient education from practitioner" (17.5%); and "low intention/motivation to adhere to a GFD (17.5%)".⁴⁵

In our study, patients with CD had an inadequate total daily energy intake. This finding was concordant with several studies from Poland, Germany and Spain.^{58–60} As for nutrients intake, among females and males, none of the study participants showed adequacy to the daily recommendations for calcium and vitamin D intake. These results, altogether, showed an unbalanced diet in terms of micronutrients, that were in accordance with those shown in several studies.^{14,61–63} On the other hand, almost all the study participants from both genders showed a high daily consumption of iron. This finding contradicts the results reported in other studies.^{60,64–66}

DS were used by 54% of the study population of which 38%, 10%, 46%, 18%, 16% and 4% used vitamin D, vitamin B12, iron, calcium, folate and probiotics, respectively. It appears that the usage of DS among individuals with CD wasn't in excess of that reported in the general population in Lebanon.⁶⁷ In our study, the most commonly used DS was vitamin D and the least used one was probiotics. According to the literature, gastrointestinal symptoms can be alleviated through prebiotics intake. It has been shown that *Bifidobacterium infantis* can alleviate gastrointestinal symptoms if administered to CD patients before meals for three weeks.⁶⁸ This can explain the occurrence and persistence of gastrointestinal manifestations among the majority of participants in the current study.

Strength and limitations

The crucial strength of this study is that it fulfills a comprehensive assessment of dietary intake and food-related patterns, biochemical parameters, body composition, and physical activity. However, this study had some limitations: 1) the selected group of patients doesn't represent the general population with CD in Lebanon; 2) it was a cross-sectional study and, thereby, our findings must be interpreted with caution; 3), physical activity levels were self-reported; 4) there is an important limitation of data on caloric, macro- and micronutrient composition of the recalled three-day record. The low sample size must also be considered.

Conclusion

Monitoring nutritional status using dietary, anthropometric, blood tests and assessing physical activity are the key components in the management of CD. The ideal GFD should be balanced with macronutrients and micronutrients, with

reasonable access and prices. Fortification/enrichment of GF foods that are frequently consumed should be encouraged to avoid deficiencies. This underlines the critical role of dietitians in education and maintenance of healthy GFD among individuals with CD.

Data availability

Underlying data

Open Science Framework: Celiac disease_Nutrition_Lebanon, <https://osf.io/qcfb3>.⁶⁹

This project contains the following underlying data:

- SPSS Celiac.sav (raw questionnaire, morphometric and serology data)

Extended data

Open Science Framework: Celiac disease_Nutrition_Lebanon, <https://osf.io/qcfb3>.⁶⁹

This project contains the following extended data:

- Questionnaire.pdf

Data are available under the terms of the [Creative Commons Zero “No rights reserved” data waiver](#) (CC0 1.0 Public domain dedication).

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Version 1

Reviewer Report 10 October 2022

<https://doi.org/10.5256/f1000research.133767.r151836>

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Anil K. Verma

Celiac Disease Research Laboratory, Department of Pediatrics, Università Politecnica delle Marche, Ancona, Italy

Yeliz Serin

Department of Nutrition and Dietetics, Faculty of Health Science, Çukurova University, Turkey

Title: Nutritional status, nutrient imbalances, food-related behaviors, and dietary supplements use among patients with celiac disease in Lebanon: a national cross-sectional study

Opinion

In this manuscript, The authors have investigated the nutritional status, food-related behaviors, caloric, macro-, and micronutrient intake and dietary supplements use among individuals with CD in Lebanon. The authors claim, there is no data related to this issue in Lebanon. On a general search on the web, I did not find any related articles from Lebanon. Hence, I appreciate the authors for paying attention to the nutritional status of celiac patients living in Lebanon. However, I have multiple doubts that I am summarising below:

Method

- **In the Selection procedure part:** In the 'exclusion criteria' the authors mentioned that they eliminated patients with malnutrition. But in Table 2, they assessed the risk of malnutrition in the patients with malnutrition. Please clarify it.
- **The line of measurements:** "*Weight and height were measured using automated scale and stadiometer. Body composition including body fat, lean body mass, and visceral fat was assessed using Bio-impedancemetry body composition analyzer (BF-511, OMRON)*" Please explain the standardization of measurements.
- **Laboratory value:** The authors mentioned the collected data from three different centers. So, I think, the normal range of parameters could be changed. How did the authors standardized the laboratory results?

- **Food-related behaviors and GFD dietary adherence:** *"Across the three-day food records, patients presenting a consumption between 3 to 10 g of gluten per day were considered as non-adherent"* In this line, please explain how you calculated gluten intake from the diet record? Which method did you use?
- **Food record calculation:** The authors showed the energy and nutrient intake in Tables 4, and 5, but they don't explain how they calculated this parameter from the diet? Which program did they use for the calculation?
- **Statistical analysis:** The authors did not mention the normality of the data set. How did they decide the normality test like (t-Test) without checking normality?
- Please also mention the full form of the SPSS first then give its abbreviation in the brackets e.g., XXX (SPSS)

Results:

- In data with a statistically significant difference, please indicate under tables that: 1) Which test did you use and 2) The cut of point of significant P value (like $p < 0,05$ or $0,001$).

Discussion:

- The discussion part contains multiple repeats of the results. On the other hand, The authors haven't interpreted the result (Like How, and why ..). For example: What are the factors that influenced adherence to GFD in Lebanon? Please after comparing the results with other studies, also give original comments related to your data.
- Please first write the whole explanation of this term "*P-E malnutrition*" (Protein-energy malnutrition)
- *"In our study, the most commonly used DS was vitamin D and the least used one was prebiotics"* The authors mentioned prebiotics, I think it was written by mistake. Please clarify this term whether probiotic or prebiotic?
- Villous Atrophy takes place in the small bowel only, no need to specify its location small bowel. Alternatively, the authors may write, partial to total villous atrophy. (First line of the main introduction)
- The first line of the Selection procedure contains results (February to July 2018). Please shift this in the results section.

Is the work clearly and accurately presented and does it cite the current literature?

Partly

Is the study design appropriate and is the work technically sound?

Partly

Are sufficient details of methods and analysis provided to allow replication by others?

Yes

If applicable, is the statistical analysis and its interpretation appropriate?

I cannot comment. A qualified statistician is required.

Are all the source data underlying the results available to ensure full reproducibility?

Yes

Are the conclusions drawn adequately supported by the results?

Yes

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: celiac disease, Gluten related disorders, HLA-DQ distribution, Non-celiac gluten sensitivity

We confirm that we have read this submission and believe that we have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however we have significant reservations, as outlined above.

Author Response 01 Nov 2022

Maha Hoteit

Dear Reviewer 3, we modified the original manuscript according to your comments:

Method

- ***"In the Selection procedure part: In the 'exclusion criteria' the authors mentioned that they eliminated patients with malnutrition. But in Table 2, they assessed the risk of malnutrition in the patients with malnutrition. Please clarify it."***
 - All the patients collected in this study were referred to us by their physicians. According to their medical files, these patients weren't malnourished. We excluded those who were diagnosed by physicians as malnourished patients because this interferes with our objectives. However, in our study, we were assessing the risk of malnutrition according to many specific criteria of which weight loss, albumin....
- ***"The line of measurements: "Weight and height were measured using automated scale and stadiometer. Body composition including body fat, lean body mass, and visceral fat was assessed using Bio-impedancemetry body composition analyzer (BF-511, OMRON)" Please explain the standardization of measurements."***
 - Percentage body fat (%BF) was measured by whole-body BIA to the nearest 0.1% using a digital scale/body composition monitor (Omron BF 511, 50 kHz, 500 μ A, Kyoto, Japan), which includes an 8-sensor technology using both hands and feet. The participants stood with bare feet on electrodes on the scale with their knees and back straight while grasping a handle that also includes electrodes with both their hands horizontally raised, elbows extended straight, and maintaining a 90°-angle to the body. A previous study that compared body composition estimates using BIA devices with DXA and whole body magnetic

resonance imaging indicated that the use of devices with additional hand electrodes provides a more accurate prediction of body composition and are suitable for public use. In order to determine percentage body fat, the device uses electrical impedance, along with the participant's height, weight, age, and gender to generate results. The readings were obtained in duplicates and the average recorded. According to the manufacturer's instructions, percentage body fat was measured two hours or more after breakfast. Omron Healthcare references for age and gender were used to classify the study participants into low, normal, high, and very high %BF.

- We followed these articles: A. Bosy-Westphal, W. Later, B. Hitze et al., "Accuracy of Bioelectrical Impedance Consumer Devices for Measurement of Body Composition in Comparison to Whole Body Magnetic Resonance Imaging and Dual X-Ray Absorptiometry," *Obesity Facts*, vol. 1, no. 6, pp. 319–324, 2008.
- Healthcare Omron, *Omron Body Composition Monitor. Instruction Manual BF 511*, Omron Healthcare Cooperation Ltd., Kyoto, Japan, 2008.
- **"Laboratory value:** *The authors mentioned the collected data from three different centers. So, I think, the normal range of parameters could be changed. How did the authors standardized the laboratory results?"*
 - The three centers were using same equipment and no issues were presented in the laboratory results.
- **"Food-related behaviors and GFD dietary adherence:** *"Across the three-day food records, patients presenting a consumption between 3 to 10 g of gluten per day were considered as non-adherent" In this line, please explain how you calculated gluten intake from the diet record? Which method did you use?"*
 - For dietary compliance, patients were accurately interviewed by a trained physician and dietitian to assess the habitual intake of serving of gluten-containing food per week (e.g. pasta, pizza, bread, or cookies). Compliance to gluten-free diet was defined as follows: very low = more than one serving of gluten containing food/week; low = up to one serving of gluten-containing food/week; good = no gluten-containing food/week. Then based on the three days record the g/d data of gluten was investigated. We followed the same methodologies adopted in the below 2 studies:
 - Ciacchi, C., Cirillo, M., Cavallaro, R., & Mazzacca, G. (2002). Long-term follow-up of celiac adults on gluten-free diet: prevalence and correlates of intestinal damage. *Digestion*, 66(3), 178–185.
<https://doi.org/10.1159/000066757>
 - Kotze L. M. (2004). Gynecologic and obstetric findings related to nutritional status and adherence to a gluten-free diet in Brazilian patients with celiac disease. *Journal of clinical gastroenterology*, 38(7), 567–574. <https://doi.org/10.1097/01.mcg.0000131720.90598.6a>

- **"Food record calculation:** The authors showed the energy and nutrient intake in Tables 4, and 5, but they don't explain how they calculated this parameter from the diet? Which program did they use for the calculation?"
 - This was already available in text. The sentence is "In addition, three-day food records were collected and analyzed using **Nutritionist Pro** software (version 3.2, AXXYA) to assess caloric, micro- and macronutrients intake. The mean value and the standard deviation (SD) were used when interpreting the nutrient's intake."
- **"Statistical analysis:** The authors did not mention the normality of the data set. How did they decide the normality test like (t-Test) without checking normality?"
 - **The normality was tested for sure.** The test used was Shapiro test.
- "Please also mention the full form of the SPSS first then give its abbreviation in the brackets e.g., XXX (SPSS)"
 - Done

Results:

- "In data with a statistically significant difference, please indicate under tables that: 1) Which test did you use and 2) The cut of point of significant P value (like $p < 0,05$ or $0,001$)."
 - Done

Discussion:

- "The discussion part contains multiple repeats of the results. On the other hand, The authors haven't interpreted the result (Like How, and why ..). For example: What are the factors that influenced adherence to GFD in Lebanon? Please after comparing the results with other studies, also give original comments related to your data."
 - Done
- "Please first write the whole explanation of this term "P-E malnutrition" (Protein-energy malnutrition)"
 - Done
- ""In our study, the most commonly used DS was vitamin D and the least used one was prebiotics" The authors mentioned prebiotics, I think it was written by mistake. Please clarify this term whether probiotic or prebiotic?"
 - *Probiotics*
- "Villous Atrophy takes place in the small bowel only, no need to specify its location small bowel. Alternatively, the authors may write, partial to total villous atrophy. (First line of the main introduction)"
 - Done
- "The first line of the Selection procedure contains results (February to July 2018). Please shift this in the results section."
 - Done

Competing Interests: No competing interests were disclosed.

Reviewer Report 05 October 2022

<https://doi.org/10.5256/f1000research.133767.r151835>

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Hamed R. Takruri

Department of Nutrition and Food Technology, Faculty of Agriculture, The University of Jordan, Amman, Amman Governorate, Jordan

Generally, the paper is good and informative, although there are some writing and structural mistakes; these are highlighted and commented on in the attached [PDF article](#). Some long sentences should be rewritten and separated into short sentences.

- It is not clear how the intake of gluten was estimated.
- It is unclear how the weight loss was evaluated; is it in reference to ideal body weight or to initial weight at the beginning of the study?

The paper can be indexed after considering the notes mentioned above.

Please find further comments in the annotated pdf hyperlinked above or at the following link:

https://f1000researchdata.s3.amazonaws.com/linked/455693.Maha_Hoteit_article_121859_-_with_referees%27_%28Hamed_Takruri%29_notes.pdf

Is the work clearly and accurately presented and does it cite the current literature?

Yes

Is the study design appropriate and is the work technically sound?

Partly

Are sufficient details of methods and analysis provided to allow replication by others?

Yes

If applicable, is the statistical analysis and its interpretation appropriate?

Yes

Are all the source data underlying the results available to ensure full reproducibility?

Yes

Are the conclusions drawn adequately supported by the results?

Partly

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Human Nutrition

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 01 Nov 2022

Maha Hoteit

Dear Reviewer 2

Thank you for your comments. We took into consideration all the raised comments and are available in the modified version. We provide below a reply point by point to your comments:

"Generally, the paper is good and informative, although there are some writing and structural mistakes; these are highlighted and commented on in the attached [PDF article](#). Some long sentences should be rewritten and separated into short sentences."

- *"It is not clear how the intake of gluten was estimated:"*
 - For dietary compliance, patients were accurately interviewed by a trained physician and dietitian to assess the habitual intake of serving of gluten-containing food per week (e.g. pasta, pizza, bread, or cookies). Compliance to gluten-free diet was defined as follows: very low = more than one serving of gluten containing food/week; low = up to one serving of gluten-containing food/week; good = no gluten-containing food/week. Then based on the three days record the g/d data of gluten was investigated. We followed the same methodologies adopted in the below 2 studies:
 - Ciacci, C., Cirillo, M., Cavallaro, R., & Mazzacca, G. (2002). Long-term follow-up of celiac adults on gluten-free diet: prevalence and correlates of intestinal damage. *Digestion*, 66(3), 178–185. <https://doi.org/10.1159/000066757>
 - Kotze L. M. (2004). Gynecologic and obstetric findings related to nutritional status and adherence to a gluten-free diet in Brazilian patients with celiac disease. *Journal of clinical gastroenterology*, 38(7), 567–574. <https://doi.org/10.1097/01.mcg.0000131720.90598.6a>
- *"It is unclear how the weight loss was evaluated; is it in reference to ideal body weight or to initial weight at the beginning of the study?"*
 - In the last 6 months prior to the interview. We followed this study : Collins N. (2003). Protein-energy malnutrition and involuntary weight loss: nutritional and pharmacological strategies to enhance wound healing. *Expert opinion on pharmacotherapy*, 4(7), 1121–1140. <https://doi.org/10.1517/14656566.4.7.1121>

"The paper can be indexed after considering the notes mentioned above."

Please find further comments in the annotated pdf hyperlinked above or at the following link:
https://f1000researchdata.s3.amazonaws.com/linked/455693.Maha_Hoteit_article_121859_-_with_referees%27_%28Hamed_Takruri%29_notes.pdf

All the comments were taken into consideration.

Competing Interests: No competing interests were disclosed.

Reviewer Report 22 July 2022

<https://doi.org/10.5256/f1000research.133767.r143011>

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Sherine Khater

Department of Gastroenterology, AP-HP, Hôpital Européen Georges Pompidou, Paris, France

It is a very interesting study about nutritional status in CD (celiac disease) patients on GFD (gluten-free diets) in Lebanon, with many parameters studied. However, I have 2 remarks:

1. Inclusion criteria include celiac disease patients attempting a GFD for at least 6 months. This should appear in the title of the article (the population is not CD patients at diagnosis but CD patients on GFD or attempting a GFD). This point should also appear in the abstract (Background section and Methods section). A very positive point was to evaluate the adherence to GFD of the patients included, which was high and thus reducing bias. In the discussion part, some references are inadequate because they studied micronutrient deficiencies in CD patients at diagnosis, and not in CD patients following a GFD (example: references 32, 34, 41). These references should be replaced by others, for example: Cardo A et al, Nutritional Imbalances in Adult Celiac Patients Following a Gluten-Free Diet, *Nutrients* 2021, 13, 2877.
2. Concerning use of probiotics in CD, there is no strong evidence to date. For this reason, I think it is better to remove this section in the discussion: *"According to the literature, gastrointestinal symptoms can be alleviated through prebiotics intake. It has been shown that Bifidobacterium infantis can alleviate gastrointestinal symptoms if administered to CD patients before meals for three weeks. This can explain the occurrence and persistence of gastrointestinal manifestations among the majority of participants in the current study"*. Persistence of symptoms might more probably be explained by associated irritable bowel syndrome, or by inadvertent gluten ingestion.

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Following a Gluten-Free Diet. *Nutrients*. 2021; **13** (8). [PubMed Abstract](#) | [Publisher Full Text](#)

Is the work clearly and accurately presented and does it cite the current literature?

Yes

Is the study design appropriate and is the work technically sound?

Yes

Are sufficient details of methods and analysis provided to allow replication by others?

Yes

If applicable, is the statistical analysis and its interpretation appropriate?

I cannot comment. A qualified statistician is required.

Are all the source data underlying the results available to ensure full reproducibility?

Yes

Are the conclusions drawn adequately supported by the results?

Yes

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Gastroenterology, celiac disease and malignant complications of celiac disease.

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 29 Jul 2022

Maha Hoteit

The authors would like to thank Reviewer 1 for reviewing the manuscript.

All comments were taken into consideration.
Changes in the manuscript are highlighted in track changes.

Competing Interests: No competing interests were disclosed.

Author Response 01 Nov 2022

Maha Hoteit

Dear Reviewer 1

All the comments raised are edited in the amended version.

Many thanks

Competing Interests: No competing interests were disclosed.

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