The relationship between Celiac Disease (CD) and obesity: A Review

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Abstract:
The relationship between obesity and Celiac Disease (CD) has been a subject of increasing interest in recent years, as both conditions continue to pose significant public health challenges globally. In people who are genetically susceptible to the condition, gluten consumption causes the autoimmune disease celiac disease, characterized by intestinal inflammation and villous atrophy. Conversely, obesity, a complex metabolic condition, is associated with excess adipose tissue accumulation and systemic inflammation. While these conditions may seem disparate, emerging research suggests a complex interplay between them.

This review aims to explore the intricate relationship between obesity and Celiac Disease, examining both potential causative mechanisms and clinical implications. Epidemiological studies have demonstrated conflicting evidence regarding the association between obesity and CD, with some indicating an increased risk among obese individuals while others report contradictory findings. Proposed mechanisms linking obesity and CD include alterations in gut microbiota composition, inflammatory cytokine profiles, and shared genetic susceptibility factors. Furthermore, the impact of obesity on CD diagnosis and management remains a topic of debate, with challenges such as underdiagnosis and delayed diagnosis in obese patients.

Understanding the relationship between obesity and Celiac Disease is crucial for optimizing patient care and public health strategies. Further research is warranted to elucidate this complex interplay’s underlying mechanisms and clinical implications, ultimately facilitating more effective prevention and management strategies for both conditions.

Keywords: Celiac disease (CD); obesity; Vit B12

Introduction:
Celiac Disease (CD) and obesity are two prevalent health issues that have garnered considerable attention due to their increasing prevalence and impact on public health worldwide. Celiac Disease is an autoimmune condition that affects the small intestine and is brought on by gluten consumption in people who are genetically predisposed leading to inflammation and damage to the intestinal villi (1). On the other hand, obesity, characterized by excessive accumulation of adipose tissue, is a multifactorial condition associated with various metabolic and inflammatory abnormalities (2).

The coexistence of these seemingly unrelated conditions has sparked interest in understanding their potential interplay. While CD is traditionally associated with malabsorption and weight loss, recent studies have suggested a more complex relationship between CD and obesity.
Understanding the nature of this relationship is essential for improving patient care and developing effective management strategies for individuals affected by both conditions.

Research investigating the association between CD and obesity has yielded conflicting findings, with some studies suggesting an increased risk of CD among obese individuals, while others report contradictory results. For example, a study by Ludvigsson et al. found a positive association between obesity in childhood and the subsequent development of CD (3). In contrast, other studies, such as that by Tata et al. did not find a significant association between body mass index (BMI) and CD risk in adulthood (4).

Proposed mechanisms underlying the relationship between CD and obesity include alterations in gut microbiota composition, dysregulation of inflammatory cytokines, and shared genetic susceptibility factors. Understanding these mechanisms is crucial for elucidating the pathophysiology of both conditions and identifying potential therapeutic targets.

Despite the growing body of research on this topic, several gaps in our understanding still need to be addressed. For instance, the impact of obesity on the clinical presentation and management of CD is not yet fully elucidated. Furthermore, the potential influence of CD on the development and progression of obesity-related complications requires further investigation.

This review aims to comprehensively examine the relationship between Celiac Disease and obesity, drawing upon existing literature to explore potential mechanisms, epidemiological associations, and clinical implications.

Celiac Disease and Obesity: A Complex Bidirectional Relationship:

Celiac disease and obesity are two prevalent health conditions with a surprisingly intricate relationship. Gluten is an autoimmune protein found in wheat, barley, and rye. It is the cause of celiac disease. Those who have celiac disease have inflammation and damage to their small intestinal lining as a result of their immune system attacking the tissue. This can cause a variety of symptoms, including diarrhea, bloating, weight loss, and fatigue. Obesity, on the other hand, is characterized by excessive body weight and is often linked to a cluster of metabolic abnormalities (5).

Recent research suggests a bidirectional link between celiac disease and obesity, highlighting the importance of considering both conditions in patient management.

The Relationship Between CD, Obesity, and Vitamin B12:

The relationship between CD, obesity, and vitamin B12 deficiency is multifaceted and influenced by various factors. Individuals with CD are at increased risk of malabsorption, including impaired absorption of vitamin B12, due to intestinal damage. Moreover, obesity may exacerbate the risk of CD-related complications and contribute to nutrient deficiencies, including vitamin B12. Conversely, vitamin B12 deficiency can lead to neurological complications, which may exacerbate symptoms in individuals with CD and obesity. Understanding these interrelationships is crucial for optimizing clinical management and improving outcomes in affected individuals (6).

Mechanisms Underlying the Association:

Several mechanisms contribute to the complex relationship between CD, obesity, and vitamin B12 status. Malabsorption resulting from intestinal damage in CD can lead to vitamin B12 deficiency, particularly in individuals with concomitant obesity. Moreover, alterations in gut microbiota composition observed in CD and obesity may further exacerbate
malabsorption and nutrient deficiencies, including vitamin B12. Additionally, obesity-related inflammation and metabolic disturbances may impact vitamin B12 metabolism and absorption, further complicating the clinical picture (6).

**Clinical Implications and Management Strategies:**

The interplay between CD, obesity, and vitamin B12 deficiency has significant clinical implications for diagnosis and management. Healthcare providers should maintain a high index of suspicion for CD in obese individuals presenting with gastrointestinal symptoms or unexplained nutrient deficiencies, including vitamin B12. Similarly, screening for obesity-related complications, such as metabolic syndrome, should be considered in individuals with CD and vice versa. Nutritional interventions, including supplementation with vitamin B12 and adherence to a balanced diet, are essential components of integrated management strategies for individuals with CD and obesity. (6).

**Impact of Celiac Disease on Obesity:**

- **Weight Loss:** Individuals with undiagnosed or untreated celiac disease often experience weight loss due to nutrient malabsorption. The damaged small intestine is unable to properly absorb essential nutrients from food, leading to deficiencies and weight loss. Studies like this one from the Mayo Clinic support this observation: (7)

- **Weight Gain:** Paradoxically, some individuals with celiac disease may experience weight gain after diagnosis and following a gluten-free diet. This can be attributed to several factors:
  - Increased Food Intake: Following a gluten-free diet, some individuals may compensate for past nutrient deficiencies by consuming more food, leading to unintended weight gain.
  - High-Calorie Gluten-Free Products: Many commercially available gluten-free processed foods can be high in calories, sugar, and fat. This can contribute to weight gain if not carefully managed within a healthy diet.
  - Reduced Physical Activity: The fear of accidental gluten exposure might lead to decreased physical activity in some individuals, hindering weight management efforts (8).

- **Increased Risk:** Research suggests a potential link between obesity and an increased risk of developing celiac disease. The exact reasons for this association are not fully understood, but several potential explanations are emerging:
  - Changes in the Gut Microbiota: It has been shown that obesity leads to modifications in the gut microbiota or the makeup of gut microorganisms. These changes to the gut flora may raise the possibility of developing celiac disease.
  - Chronic Inflammation: Obesity is a state of chronic, low-grade inflammation. This chronic inflammation might play a role in the initiation or progression of celiac disease.
  - Dietary Factors: A diet high in refined carbohydrates and gluten-rich foods may be associated with an increased risk of celiac disease, although more research is needed to confirm this association (9).
- Diagnostic Challenges: The atypical presentation of celiac disease in obese individuals can pose diagnostic challenges. Symptoms like bloating, fatigue, and even mild diarrhea may be attributed to obesity itself, delaying diagnosis and proper treatment. Additionally, studies suggest that some individuals with celiac disease and obesity may have less severe intestinal damage, potentially leading to milder or atypical symptoms.

Research published by the National Institutes of Health explores the challenges of diagnosing celiac disease in obese adults: (10).

**Treatment and Management:**

- Gluten-Free Diet: Promotes weight normalization in individuals with celiac disease [7]. However, as discussed previously, weight management for individuals with celiac disease can be complex.

- Weight Management Strategies: For individuals with celiac disease and obesity, a comprehensive weight management plan is crucial. This plan should be tailored to the individual's needs and may include:
  - Healthy Eating: A balanced diet rich in fruits, vegetables, whole grains (when tolerated), and lean protein is essential for weight loss and overall health. Gluten-free alternatives should be chosen wisely, focusing on naturally gluten-free options like fruits, vegetables, legumes, and lean protein sources (11). Opting for whole grains whenever possible can provide additional fiber and nutrients (12).
  - Regular Exercise: Engaging in regular physical activity is a cornerstone of weight management. Aim for at least 30 minutes of moderate intensity exercise most days of the week (13). Exercise helps to burn calories, improve insulin sensitivity, and promote overall health.

  - Behavioral Changes: Implementing lifestyle changes like mindful eating and stress management can be highly beneficial for weight management (14).

  - Mindful eating practices can help individuals with celiac disease develop a healthy relationship with food and avoid emotional eating (15). Additionally, managing stress levels can be crucial, as stress can sometimes trigger unhealthy eating habits (15).

**Additional Considerations:**

Nutritional Counselling: Consulting with a registered dietitian experienced in celiac disease can be invaluable for creating a personalized gluten-free meal plan that supports both weight management and nutrient needs (17).

**Conclusion:**

The relationship between celiac disease (CD) and obesity is undeniably complex, characterized by a myriad of interconnected factors that influence disease development, progression, and management. This intricate interplay challenges traditional views of these conditions as distinct entities and underscores the importance of adopting a holistic approach to patient care.

In conclusion, the bidirectional relationship between CD and obesity highlights the need for a paradigm shift in both clinical practice and research. Recognizing the overlapping pathophysiological mechanisms and shared risk factors is essential for optimizing patient outcomes and developing integrated management strategies.

As elucidated through this review, individuals with CD are not immune to the challenges posed by
obesity, and vice versa. CD can manifest in obese individuals with atypical symptoms, leading to underdiagnosis and delayed treatment. Conversely, obesity may exacerbate the risk of CD-related complications and hinder effective management strategies.

Moreover, the bidirectional relationship between CD and obesity extends beyond the realm of clinical manifestations to encompass socioeconomic, psychological, and public health implications. Addressing the complex interplay between these conditions requires a multidisciplinary approach involving gastroenterologists, dietitians, endocrinologists, mental health professionals, and policymakers.

Moving forward, further research is warranted to unravel the underlying mechanisms driving the relationship between CD and obesity and identify novel therapeutic targets. Longitudinal studies exploring the impact of weight management interventions on CD outcomes and vice versa are needed to inform evidence-based clinical guidelines.

In essence, the relationship between CD and obesity serves as a poignant reminder of the interconnectedness of various health conditions and the importance of adopting a comprehensive and patient-centered approach to healthcare. By recognizing and addressing this complex interplay, healthcare providers can improve diagnostic accuracy, optimize treatment strategies, and ultimately enhance the quality of life for individuals affected by these conditions.

**Conflict of interest**

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