#### REVIEW

## Anxiety and Depression in Eosinophilic Esophagitis: A Scoping Review and Recommendations for Future Research

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**Abstract:** Eosinophilic esophagitis (EoE) is a rapidly emerging chronic disease with limited treatments. Impacting both children and adults, EoE and its treatments have the potential to significantly reduce psychological functioning, including increasing anxiety and depression. In this paper, we provide a scoping review of the scientific literature on anxiety and depression in both pediatric and adult patients. We aim to document the current state of research, identify knowledge gaps, recognize unique challenges that EoE patients may face as they relate to psychosocial function, and offer suggestions for future research directions. Based on the current review, both adult and pediatric patients with EoE are at risk for anxiety and depression symptoms, both due to the disease process and the prevailing dietary treatments. Significant gaps exist in the current understanding of anxiety and depression in EoE, providing ample opportunity for future studies to address this important issue.

#### Keywords: eosinophilic esophagitis, anxiety, depression, quality of life, scoping review

#### Introduction

Eosinophilic gastrointestinal disorders (EGIDs) are immune-mediated chronic diseases with the potential to impact the entirety of the digestive system.<sup>1</sup> The most commonly diagnosed EGID, eosinophilic esophagitis (EoE), is characterized by chronic infiltration of the esophagus by eosinophils, normally absent in esophageal tissues. The diagnostic cutoff for EoE is greater than 15 eosinophils per high power field retrieved by biopsy. The prevailing theory is eosinophilic infiltration is due to exposure to environmental antigens, typically in foods, that triggers the allergic response. Structural changes may occur, including the presence of esophageal rings, furrows, exudates, and stricture.<sup>2</sup> The most common symptom of EoE in adult patients is dysphagia, or difficulty swallowing, that may lead to food impaction. Other symptoms include heartburn and chest pain, which are the most common symptoms in children along with failure to thrive and poor appetite. Treatments for EoE are limited to swallowed, topical corticosteroids and elimination diets.<sup>2,3</sup>

In addition to EoE, other EGIDs include eosinophilic gastritis (EG; impacting the stomach), eosinophilic gastroenteritis (EGE; impacting the stomach and small bowel) and eosinophilic colitis (EC; impacting the colon).<sup>4</sup> These EGIDs are considered much rarer than EoE at the present time, and symptoms include nausea, vomiting, loss of appetite, abdominal pain, and diarrhea. Treatments are also limited to corticosteroids and dietary modification.

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© 2019 Taft et al. This work is published and licensed by Dove Medical Press Limited. The full terms of this license are available at https://www.dovepress.com/terms.php you hereby accept the Terms. Non-commercial uses of the work are permitted without any further permission from Dove Medical Press Limited, provided the work is properly attributed. For permission for commercial use of this work, please see paragraphs A2 and S of our Terms (https://www.dovepress.com/terms.php). Currently, EoE is estimated to impact 10 to 57 per 100,000 people globally, with increasing incidence and prevalence rates.<sup>5</sup> The estimated cost of EoE approaches \$1 billion dollars annually, derived from outpatient visits, endoscopies, hospitalizations including emergency room visits, and medications.<sup>6</sup> Additional costs are attributed to the need to follow special elimination diets, requiring patients to shop at specialty stores, or the use of expensive elemental formulas. Many patients are diagnosed with an EoE as a child, further adding to the extended burden of this disease.

As EoE has gained more attention in gastroenterology practice and research, the evaluation of their impact on mental well-being and health-related quality of life has also increased. However, significant gaps remain in the literature in our understanding of the relationships between EoE and these constructs, especially compared to other chronic digestive diseases such as inflammatory bowel disease<sup>7,8</sup> and irritable bowel syndrome.<sup>9</sup> As such, we aim to review the existing literature as it relates to two specific mental health concerns in patients with EoE: anxiety and depression, in both pediatric and adult patient populations. We also aim to provide suggestions for future lines of inquiry in anxiety and depression in EoE, as well as other potential aspects of mental health that may be impacted by these diseases.

#### **Methods**

A scoping review of studies published through July 2019 (no start date) in which participants had a diagnosis of at least one EGID type was performed via the online databases PubMed, PsycINFO, and Google Scholar following PRISMA guidelines (Figure 1).

## Inclusion Criteria

We included all studies that met the following criteria: a) published in English in a peer-reviewed journal, b) participants were diagnosed with at least one EGID, and c) anxiety and/or depression was evaluated using standardized assessment tools. Studies that measured health-related quality of life were evaluated and included if anxiety or depression was part of the HRQOL assessment tool utilized.

## Procedures

Articles were identified and stored in an EndNote database. Titles or abstracts were returned via the following keyword search combinations: eosinophilic gastrointestinal disorders, eosinophilic esophagitis, eosinophilic gastritis, eosinophilic gastroenteritis, eosinophilic colitis, anxiety, depression, psychosocial, psychological, mental health. Each unique abstract was examined by two raters (TT, LG) for inclusion criteria and any that were explicitly unmet (e.g., study of nonhumans, no mention of anxiety or depression) were discarded. Inter-rater agreement was 100%. Reference lists of identified articles and book chapters were also reviewed for additional studies. Unpublished manuscripts, case reports, and dissertations are not included.

The majority of studies were published in the United States and Switzerland. Both adult and child/adolescent studies are included in this review and the findings are organized adult and pediatric studies. To date, no research evaluates anxiety and depression in non-EoE EGIDs. As such, this review is limited to EoE.

## Adult Patients with EoE General Findings

The literature search yielded only a handful of studies directly measuring anxiety or depression in adult EoE patients. Often times, anxiety and depression are included in studies evaluating other constructs such as HRQOL, social interactions, and disease stigmatization. As such, reviews of these patient outcomes are included within the context of how they are associated with anxiety and depression.

## Anxiety and Depression

Few studies have evaluated the prevalence and impact of anxiety or depression in adult patients with EoE (Tables 1 and 2). A 2019 study sought to characterize a cohort of adult EoE patients through a retrospective medical record review, which included assessing the prevalence of several psychiatric comorbidities using ICD-9 and ICD-10 codes.<sup>10</sup> Overall, 31% of patients had at least one psychiatric or neuropsychiatric comorbidity, with 12% of the sample demonstrating a diagnosis of depression, followed by anxiety, which was reported by 9.3% of patients. Further, the authors found these rates were significantly higher than the general hospital population, although comparable to the natural history of each respective disorder.<sup>10</sup> Another study applied the Hospital Anxiety and Depression Scale-8 (HADS-8) to a cohort of Spanish patients with EoE. Results demonstrated that 31.1% and 9.8% reported anxiety and depression in the "probable-certain" range (score > 8), respectively.<sup>11</sup> These



Figure I PRISMA diagram for selection of sources of evidence.

findings remained relatively stable at the 12-week re-test period.

Research suggests potential sex differences in adult patients with EoE, specifically with women reporting increased symptoms of anxiety and depression compared to men.<sup>12</sup> These findings are consistent with research in the general population.<sup>13–15</sup> However, more research is warranted before conclusive statements can be made.

#### Illness Stigma

Illness stigma may also impact anxiety and depression in adult patients with EoE. Greater internalized and perceived illness stigma was found to be significantly associated with increased anxiety and depression symptoms.<sup>12,16</sup> Further, perceived and internalized stigma were each significant predictors of both anxiety and depression, even when controlling for variables such as disease severity, duration, and gender.<sup>12,16</sup>

## Psychological Impact of EoE Treatment

There are several options for EoE treatment, including pharmacological therapy, procedural intervention (e.g. balloon distention), and dietary modification, all of which have the potential of impacting an individual's psychosocial functioning. In terms of medication, patients with EoE may report fear or worry about potential side effects,<sup>17</sup> which can exacerbate an already stressful disease experience. Patients using dietary therapy may also experience negative impacts on their psychosocial functioning, in some cases even more so than those exclusively using medications.<sup>18</sup> Dietary therapy requires adherence, which can be burdensome and inconvenient,<sup>17,19</sup> and has been

	Author Group	Year	Study Design	Sample Size	Assessments Used	Estimates Obtained	
I	Leigh and Spergel <sup>10</sup>	2019	Retrospective Chart Review	34	ICD-9/ICD-10 code	12% Had diagnosis of Depression.	
2	Wang et al <sup>19</sup>	2018	Cross-sectional	42	NIH-PROMIS Depression Scale	Depression scores higher in former users of SFED compared to current users.	
3	Taft et al <sup>26</sup>	2011	Scale Development	201	EoE-QOL-A	Emotional Impact is a subscale on EoE- specific measure of QOL.	
4	Lucendo et al <sup>11</sup>	2014	Cross-sectional	170	Hospital Anxiety & Depression Scale	9.8% Scored above cutoff for depression.	
5	Guadagnoli and Taft <sup>12</sup>	2019	Cross-sectional	149	NIH-PROMIS Depression Scale	Stigma internalization is positively correlated with increased depression.	
6	Guadagnoli et al <sup>16</sup>	2017	Cross-sectional	149	NIH-PROMIS Depression Scale	Stigma perception is positively correlated with increased depression.	

Table I Relevant Studies Examining Depression in Adults with EoE

found to negatively impact the emotional aspects of HRQOL in adult patients with EoE.<sup>20</sup> Further, research has found that diet-related anxiety, such as worry about food being unavailable and time required to prepare foods, is associated with decreased dietary adherence for adult patients on the six-food elimination diet.<sup>19</sup> In addition, anxiety and depression levels appear to be higher in patients identified as former users of the six-food elimination diet, suggesting a relationship between emotional distress and dietary use, although directionality cannot be concluded.<sup>19</sup>

## Social Impacts of EoE

Finally, EoE may impact patients' social experience, which can in turn, affect emotional well-being. Patients with EoE report experiencing social embarrassment or distress due to their diagnosis, including negative experiences dining out or attending work interviews, worry about having a choking episode in public, and concern about having to hide the attack from others.<sup>17</sup> Embarrassment and shame may contribute to perceived or internalized illness stigma, which can also negatively affect anxiety and depression symptoms in patients with EoE.<sup>12,16</sup>

Interestingly, despite these findings, patients typically endorse feeling open to disclosing their illness to friends and family.<sup>17</sup> The benefits of social support have been studied in a variety of chronic illnesses and are established as an important protective factor for prevention of adverse mental health outcomes and promotion of positive health behaviors.<sup>21,22</sup> Thus, encouraging discussion about EoE and cultivating support from friends and family may aid as a buffer the potentially harmful outcomes.

## Relationship with HRQOL

While literature on anxiety and depression in adult patients with EoE is scarce, there has been progress in the field evaluating the impact of EoE on health-related quality of life (HRQOL). Research has consistently found that EoE patients experience reduced HRQOL, due to several facets of their disease, which has resulted in psychological and social implications.<sup>18,20,23,24</sup> Notably, adult patients with EoE view HRQOL as one of the most important outcomes in long and short-term therapy, even when compared to reduction in biological and histological activity,<sup>25</sup> underscoring HRQOL as an important facet of the overall EoE patient experience.

The demand for a better understanding of the impact of EoE on psychosocial functioning was augmented by the development of the Adult Eosinophilic Esophagitis Quality of Life (EoE-QOL-A) questionnaire, a tool to assess HRQOL specific to the EoE population.<sup>26</sup> In addition to a total score, the EoE-QOL-A is comprised of several subscales, including emotional impact, social impact, eating/diet impact, choking anxiety, and disease anxiety, all of which have individual subscale scores, which allows for identification of specific underlying issues patients may be experiencing.

The emotional impact and choking and disease anxiety subscales are of particular interest, as it evaluates aspects

	Author Group	Year	Study Design	Sample Size	Assessments Used	Estimates Obtained	
I	Wang et al <sup>19</sup>	2018	Cross-sectional	42	NIH-PROMIS Anxiety Scale	Diet-related anxiety may impact SFED adherence; Anxiety higher in former users of SFED compared to current users.	
2	Stern et al <sup>18</sup>	2018	Prospective	167	EoE-QOL-A	Anxiety (measured under QOL), is associated with symptom severity but not endoscopy or histology results. Prior food impaction influences choking anxiety.	
3	Safroneeva et al <sup>2</sup>	2018	Cross-sectional	109	EoE-QOL-A	Anxiety (measured under QOL) is associated with patient treatment goals.	
4	Lucendo et al <sup>20</sup>	2018	Cross-sectional	170	EoE-QOL-A	Disease anxiety and choking anxiety were the highest scoring domains of QOL.	
5	Taft et al <sup>17</sup>	2011	Qualitative	24	Semi-Structured Interview	Disease anxiety and choking anxiety are important aspects of QOL.	
6	Taft et al <sup>26</sup>	2011	Scale development	201	EoE-QOL-A	Disease anxiety and choking anxiety are 2 subscales on EoE-specific measure of QOL.	
7	Lucendo et al <sup>11</sup>	2014	Prospective	170	Hospital Anxiety and Depression Scale	31.1% of patients scored above the cutoff for anxiety.	
8	Guadagnoli and Taft <sup>12</sup>	2019	Cross-sectional	149	NIH-PROMIS Anxiety Scale	Stigma internalization is positively correlated with increased anxiety.	
9	Guadagnoli et al <sup>16</sup>	2017	Cross-sectional	149	NIH-PROMIS Anxiety Scale	Stigma perception is positively correlated with increased anxiety.	
10	Leigh and Spergel <sup>10</sup>	2019	Retrospective chart review	34	ICD-9/ICD-10	9.3% Had diagnosis of anxiety.	
11	Safroneeva et al <sup>48</sup>	2018	Cross-sectional	98	EoE-QOL-A	Symptoms severity was correlated with HRQOL scale and predicted HRQOL scores, including anxiety.	

Table 2 Relevant Studies Examining Anxiety in Adults with EoE

of anxiety and depression within HRQOL. The emotional impact subscale assesses the degree to which patients have felt like life is less enjoyable or felt helpless or isolated because of EoE, and was found to have the greatest influence on decreased HRQOL in a sample of patients with EoE in a recent study.<sup>18</sup>

The disease anxiety and choking anxiety subscales are also important contributors to HRQOL in adult EoE patients.<sup>17</sup> In fact, a separate study found that disease anxiety and choking anxiety had the largest impacts on overall HRQOL in the EoE population.<sup>11,20</sup> EoE symptoms, such as dysphagia or food impaction, can produce a significant amount of fear and concern for patients,<sup>17,18</sup> and increased symptom severity is a significant predictor of both disease and choking anxiety.<sup>27</sup> Further underscoring the association between emotional distress and HRQOL is the finding that increased scores on more general measures of anxiety and depression are significantly associated with decreases in HRQOL (as measured by the EoE-QOL-A).<sup>11,26</sup> However, some research suggests the correlation between anxiety and HRQOL is stronger than the relationship between HRQOL and depression.<sup>11</sup>

# Limitations and Future Research for Adults with EoE

There is limited understanding of the role of anxiety and depression in adult patients with EoE. To our knowledge, there is one qualitative study evaluating these constructs, and while there is an increasing number of quantitative studies, the research is generally lacking in rigor and efficacy.

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No prospective studies exist that are specifically aimed to evaluate clinical anxiety and/or depression in patients with EoE. In terms of prevalence, one study exists reporting the prevalence of anxiety and depression diagnoses in the adult EoE population.<sup>10</sup> While this helpful on a broad level, given the nature of retrospective reviews, we cannot confirm the accuracy of the diagnostic codes. For example, patients may be in remission for their psychiatric disorder but still have an active ICD code, or diagnoses may have been determined without appropriate psycho-diagnostic evaluation. Thus, results should be interpreted with caution.

A few studies applied self-reported measures of anxiety and depression, such as the PROMIS Anxiety, PROMIS Depression, and Hospital Anxiety and Depression Scale (HADS). These scales are particularly useful in screening to understand the magnitude and direction of anxious or depressive symptoms. However, they are not sufficient in diagnosing a clinical level of anxiety or depression, which would require an additional evaluation by a certified mental health professional.<sup>28</sup> In addition, several of these studies included anxiety and depression as secondary outcomes, resulting in lack of emphasis on the findings. Thus, future research should prioritize evaluating anxiety and/or depression as main outcomes, as further investigate the relationship between these constructs and other relevant clinical and psychosocial information.

The overwhelming majority of research in adult patients with EoE uses HRQOL scales (i.e. EoE -QOL -A) that include anxiety or emotional-related subscales, to assess for anxiety and depression symptoms. These studies capture the impact of EoE-specific anxiety and emotional distress, as compared to anxiety and depression as a general mental health concern (such as in the studies discussed above), which are viewed and assessed as different constructs. For example, a patient may not endorse general anxiety and worry in his or her everyday life but may experience disease or choking anxiety. Both constructs are important to consider in this population and could possibly produce different outcomes. While clinical anxiety and depression may result in an enhanced characterization of the EoE population, EoE-specific anxiety and depression can identify the more nuanced and potentially clinically relevant associations. Thus, future research should continue to pursue EoE-specific distress, such as emotional impacts as well as disease and choking anxiety, and the influence it has on adult patients with EoE.

## Children and Adolescent Patients with EoE

## **General Findings**

As the prevalence of EoE appears to be quickly growing,<sup>29,30</sup> a better understanding of the impact of EoE on quality of life in children becomes more critical. The existing literature establishes a link between EoE and poorer HRQOL, with symptoms and disease status directly correlating with lower HRQOL.<sup>31,32</sup> Within the larger body of research on HRQOL in EoE, evidence points to greater emotional and behavioral problems and worsened adjustment as children get older.<sup>33</sup> Based on the scant studies of emotional symptoms within HRQOL, disease-specific symptoms such as epigastric pain and certain treatment options (e.g., food restrictions and g-tubes) may be most predictive of emotional symptoms.<sup>33,34</sup>

In examining anxiety and depression specifically, however, the research is limited. Many studies of pediatric health-related quality of life collapse emotional symptoms on the PedsQL measure into one "psychosocial functioning" domain combined with school and social functioning, or include "worry" measured within the EoE module of the PedsQL.<sup>31,35</sup>

#### Anxiety

Studies examining anxiety specifically suggest that youth with EoE do have higher rates of anxiety symptoms compared to healthy populations<sup>29,33</sup> (Table 3). In the study of 705 patients, 15.5% reported anxiety and depression, with 19% of older patients (11-17 years) endorsing these symptoms compared to 9.3% of younger patients (<11 years).<sup>29</sup> In a smaller cohort of 64 patients, reported rates of anxiety were 41% among children, but not stratified by age.<sup>33</sup> These higher rates may be linked with symptoms and treatments unique to EoE, including worries about symptoms,<sup>36,37</sup> dietary restrictions, and g-tubes.<sup>34,38</sup> In the review by Klinnert et al clinical observations found patients with increased anxiety related to needing a G-tube and in those with comorbid food allergies which heightened fears about a lifethreatening reaction. Qualitative interviews corroborate these concerns with anxieties traversing food reactions and choices, and having endoscopies.<sup>36</sup> The common intervention of dietary restriction likely increases worry about foods (by children and their caregivers), as well as fears of embarrassment in social situations.34,39 The specific symptoms most often identified include worry about disease and symptoms, physical symptoms of anxiety and autonomic arousal,<sup>40,41</sup> and school avoidance.<sup>41</sup> It should be noted that

	Author Group	Year	Study Design	Sample Size	Assessments Used	Estimates Obtained	
I	Jose et al <sup>41</sup>	2016	Cross-sectional (Abstract)	8	Screen for Child Anxiety- Related Disorders	38% Met criteria for anxiety.	
2	Harris et al <sup>33</sup>	2013	Retrospective chartrReview	64	I-hr Pediatric Behavioral Health Assessment	41% Reported anxiety, anxiety symptoms increased with age.	
3	Franciosi et al <sup>36</sup>	2012	Qualitative	42	Semi-Structured Interview	Worry about symptoms and EoE were specific themes.	
4	Cortina et al <sup>40</sup>	2010	Cross-sectional	108	Multidimensional Anxiety Scale for Children	Compared to healthy controls, children and adolescents with EOE reported more physical symptoms of anxiety and autonomic arousal.	
5	Case et al <sup>38</sup>	2017	Cross-sectional	46 families	Revised Children's Anxiety and Depression Scale, Pediatric Quality of Life Scale for EoE	50% of participants reported high frequency worry related to special diets, measured by PedsQL-EoE Feelings subscale.	
6	Chehade et al <sup>29</sup>	2018	Prospective	705	Self-Reported History of Anxiety or Depression	24.0% in subjects ≥18 years of age, 9.0% in those 11–17 years of age, 9.3% in children <11 years of age.	
7	Jose et al <sup>41</sup>	2017	Cross-sectional	20	Screen for Child Anxiety- Related Disorders	Caregivers of children with EoE report more symptoms of anxiety disorder, panic disorder, and school avoidance; compared to general population.	
8	Lynch et al <sup>35</sup>	2018	Review		N/A	Higher rates of anxious symptoms in youth with EoE are associated with sleep problems and pain, all of which impact HRQOL (no study has yet examined these four constructs at once).	
9	Lynch et al <sup>43</sup>	2017	Cross-sectional	50	Pediatric Quality of Life Scale for EoE	Caregiver reports rated children with EoE and persistent pain as having more anxiety symptoms compared to controls.	

Table 3 Relevant Studies Examining Anxiety in Children/Adolescents with EoE

a majority of studies including anxiety as a variable discuss anxiety at the symptom level rather than representing actual anxiety disorders. Some evidence shows that anxiety symptoms increase with age,<sup>33</sup> suggesting the importance of developmental considerations.

#### Depression

Although anxiety has received more attention than depression in studying emotional symptoms in children and adolescents with EoE, results of studies to date suggest that mood is an important area in which to increase understanding (Table 4). When measured with anxiety symptoms, some evidence shows depression may also increase with age at higher rates than the general population.<sup>29</sup>

Depressive symptoms have been found to contribute to medical non-adherence in youth with EoE,<sup>42</sup> and to be related to persistent epigastric pain and greater sleep disturbance.<sup>43</sup> In one study, children and adolescents with EoE reported more symptoms on subscales of anhedonia, negative mood, and negative self-esteem compared to healthy controls.<sup>40</sup> This early evidence base justifies depression and depressive symptoms as an important component of studying HRQOL in pediatric EoE.

## Limitations and Suggestions for Future Research for Children with EoE

Data from the limited research of anxiety and depression in pediatric EoE populations indicate this area needs

	Author Group	Year	Study Design	Sample Size	Assessments Used	Estimates Obtained	
I	Chehade et al <sup>29</sup>	2018	Cross-sectional	705	Self-Reported History of Anxiety or Depression	24.0% in subjects $\geq$ 18 years of age, 9.0% in those 11–17 years of age, 9.3% in children <11 years of age	
2	Hommel et al <sup>42</sup>	2012	Cross-sectional	96	Behavior Assessment System for Children, 2nd Edition	Depressed patients being more likely (OR=7.27) to be non-adherent than non- depressed patients	
3	Lynch et al <sup>35</sup>	2018	Cross-sectional	50	Pediatric Quality of Life Scale for EoE	Caregivers of children with EoE and persistent pain reported significantly more depressive symptoms in their children compared to healthy controls	
4	Case et al <sup>38</sup>	2017	Cross-sectional	20	Revised Children's Anxiety and Depression Scale, Pediatric Quality of Life Scale for EoE	50% of participants reported high frequency anger and sadness related to special diets, measured by PedsQL-EoE Feelings subscale	
5	Cortina et al <sup>40</sup>	2010	Cross-sectional	108 (Mothers & Children)	Children's Depression Inventory	Children and adolescents with EoE reported greater overall depressive symptoms compared to controls, and more symptoms on three subscales: anhedonia, negative mood, negative self-esteem	
6	Lynch et al <sup>43</sup>	2018	Review			Higher rates of depressive symptoms in youth with EoE are associated with sleep problems and pain, all of which impact HRQOL (no study has yet examined these four constructs at once)	

Table 4 Relevant Studies Examining Depression in Children/Adolescents with EoE

further study to provide better understanding of the role of psychological symptoms across the developmental continuum of life with EoE, and to inform more holistic treatment interventions. Future research should more systematically not only include depression and anxiety as constructs, but should develop a more consistent approach to conceptualization and measurement.

Associations between factors such as internalizing symptoms, sleep disturbance, and pain warrant more analysis. An improved understanding of how these symptoms interrelate would inform more effective clinical interventions, including behavioral health treatments potentially targeting pain and sleep in order to decrease internalizing symptoms.

Finally, caregiver anxiety has been shown to be elevated, especially for those of younger children, due to worries about growth, nutrition, and avoiding harmful foods.<sup>34</sup> As parent anxiety is known to increase the risk of child anxiety, this potential association also deserves more research in pediatric EoE. Screening for caregiver anxiety and providing appropriate referrals such as behavioral health interventions for caregivers could be an important part of treatment for pediatric EoE.

## Conclusions

This scoping review of anxiety and depression studies in pediatric and adult patients with EoE demonstrates a significant gap in the existing literature and the necessity of more research to understand the true impacts of these mental health co-morbidities on patient outcomes. However, we can conclude that EoE does increase the risk of developing symptoms of anxiety and depression, which in turn may reduce HRQOL and treatment adherence. The lack of treatment options may be a driver of anxiety, especially in those patients who need to follow strict elimination diets indefinitely. Impacts on eating and food-specific anxieties are legitimate concerns that treating physicians and dietitians should actively monitor, including screening for restrictive behaviors beyond what would be considered reasonable (i.e. Avoidant Restrictive

Measure Title	Anxiety	Depression	Number of Items				
Adults 18 years+							
Hospital Anxiety and Depression Scale (HADS) Generalized Anxiety Disorder – 7 (GAD-7) Patient Health Questionnaire – 9 (PHO-9)	✓ ✓	✓ ✓	8 7 9				
NIH-PROMIS Mental Health – Anxiety NIH-PROMIS Mental Heath – Depression Kessler 10	✓ ✓	J J	7 8 14				
Children 17 years or Younger			•				
Patient Health Questionnaire for Adolescents (PHQ-A) Mood & Feelings Questionnaire – Short Form (MFQ-SF) Childhood Anxiety Sensitivity Index (ASI) Screen for Child Anxiety-Related Disorders (SCARED) <sup>a</sup>	1	✓ ✓ ✓	9  3  6 4				

Note: <sup>a</sup>Completed by parent.

Food Intake Disorder (ARFID)).<sup>44</sup> ARFID is a condition defined by the Diagnostic and Statistical Manual for Mental Disorders – Fifth dition,generally applied to feeding issues in children. Recently ARFID specific to coping with digestive diseases has become a growing area of attention, including a 2019 case report of two children with EoE.<sup>45</sup>

We recommend gastroenterologists who treat EoE, and other EGIDs for that matter, conduct routine screens for anxiety and depression. There are several brief measures easily implemented in clinical practice such as the PROMIS mental health scales, the PHQ-9, and the GAD-7 (Table 5). In addition to screening, referral lines to mental health professionals specializing in health psychology, behavioral medicine, or psycho-gastroenterology should be established so patients can be provided with resources once anxiety or depression symptoms are identified. Effective psychological interventions for anxiety and depression, such as cognitive behavioral therapy or acceptance and commitment therapy, while not formally evaluated in EoE, have been tested in other chronic digestive diseases and demonstrate sufficient efficacy.<sup>46,47</sup>

#### Disclosure

The authors report no conflicts of interest in this work.

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