

# Nutrition Assessment, Interventions, and Monitoring for Patients with Celiac Disease: An Evidence Analysis Center Scoping Review



#### ABSTRACT

The objectives of this scoping review were to identify and characterize studies examining nutrition assessment, interventions, and measures to monitor gluten-free diet (GFD) adherence/compliance in patients with celiac disease (CD). An electronic literature search of four databases (Cochrane Database for systematic reviews, CINAHL, Embase, and Ovid MEDLINE) was conducted to identify articles examining nutrition care in CD individuals. Except for narrative review, grey literature, and case study/report, all types of peer-reviewed articles published between January 2007 and August 2018 were eligible. There were a total of 10,823 records; 10,368 were excluded during the first round of screening due to irrelevancy and/or duplication. Of the 455 full-text articles that were assessed, 292 met the criteria and were included. Most of the studies were observational studies (n=212), followed by experimental trials (n=50), evidencebased practice guideline (EBPG)/report/statement (n=16), and systematic review (SR) (n=14). Nine original studies examined assessment, focusing mainly on different tools/ways to assess GFD adherence. The majority of the included original articles (n=235) were in the nutrition intervention category with GFD, oats, and prebiotics/probiotics as the top-three most studied interventions. There were eight SRs on GFD and five on oats. One SR and 21 original studies investigated the effectiveness of different measures to monitor GFD adherence/compliance. Although recent CD EBPGs were identified, different methods with varying levels of rigor, in terms of literature search and assessment of evidence strength, were used. Based on this scoping review, interventions focused on gluten-free diet and oats have been significantly covered by either SRs or EBPGs. Studies related to prebiotics/probiotics and education program/counseling focused interventions, as well as assessment, in CD patients have increased in recent years. Thus, it might be beneficial to conduct SRs/ EBPGs focused on these topics to guide practitioners. J Acad Nutr Diet. 2020;120(8):1381-1406.

ELIAC DISEASE (CD) IS defined as "a small intestinal immune-mediated enteropa- thy" triggered by gluten ingesgeneticallv tion in predisposed individuals.<sup>1</sup> Serologic tests (eg, antitissue transglutaminase antibodies, endomysial antibodies, and deamidated gliadin peptide antibodies), intestinal biopsies, and sometimes genetics tests are used as part of the diagnostic process to confirm CD.<sup>2</sup> This condition affects approximately 1% of the population, with slight prevalence differences among countries.<sup>3</sup> Currently, the most effective treatment for CD is a lifelong gluten-free diet.<sup>4,5</sup> Among those with this disease, higher compliance to a gluten-free diet is associated with better health outcomes (eg, returning to normal growth/development in children and fewer complications).<sup>2</sup> However, adhering to this

2212-2672/Copyright © 2020 by the Academy of Nutrition and Dietetics. https://doi.org/10.1016/j.jand.2019.09.019 Available online 14 January 2020 type of restrictive diet is often challenging, and it also can contribute to potential nutrition imbalance (eg, micronutrients deficiency).<sup>6</sup> Therefore, nutrition care is crucial for individuals with CD and registered dietitian nutritionists can play an important role at each step of the Nutrition Care Process from nutrition assessment to nutrition monitoring and evaluation to improve the health of those patients.

In 2009, the Evidence Analysis Library (EAL) at the Academy of Nutrition and Dietetics published a guideline on CD. It included recommendations for nutrition assessment, nutrition intervention, and nutrition monitoring and evaluation-all based on systematic reviews.<sup>7</sup> Because the last guideline was 10 years ago, the EAL set out to update its current guideline to incorporate any new evidence from the past 10 years. Thus, the first step of the process is to conduct a scoping review to investigate and map out the availability of new literature. Similar to a systematic review, a scoping review follows the same methodological rigor

(eg, performing a comprehensive literature search in various databases); the only difference is that a scoping review does not evaluate the methodological quality of the included studies.<sup>8</sup> That is because the purpose of a scoping review is for researchers to evaluate whether or not there is enough evidence (or in which area) to undertake systematic reviews and/or evidencebased practice guidelines, and also to see whether there are recent systematic reviews or guidelines with similar scope and methodological rigor that could potentially be adapted or recommended.8

Therefore, the aim of this scoping review was to identify and characterize studies examining the validity and reliability of nutrition assessment methods, nutrition interventions, and tools/measures to monitor gluten-free diet adherence/compliance among individuals with CD. This resulted in three research questions for this scoping review:

1. In individuals with CD, what is the availability of the literature

### Cochrane Database for systematic reviews [August 29, 2018]

#### #1 celiac\* or coeliac\*

#2 (toxicity or inulin or (Photon Absorptiometry or dexa) or skinfold\* or (body next composition) or (arsenic\* or heavy metal\*) or (nutrition\* or diet\* or feed\* or eating or malnutrition\*) or (probiotic\* or prebiotic\* or microbial supplement or vitamin\* or micronutrient or B vitamins or folate or niacin or B12 or riboflavin or calcium or iron or zinc or magnesium or fiber or protein\* or grain\* or seed\* or starch\* or cereal or gliadin\* or glutenin\*) or (gluten\* or oat or Avena or wheat or Triticum or rye or Secale or barley or Hordeum or triticale or kamut or spelt or semolina or durum) )

#3 #1 and #2

#4 accession near2 pubmed

#5 accession near3 embase

#6 #3 not (#4 or #5)

Hits: 97

### CINAHL (ebsco)

(MH "Celiac Disease+") OR TI ( (celiac\* or coeliac\*) ) OR AB ( (celiac\* or coeliac\*) )

#### AND

( (MH "Gluten") OR (MH "Diet, Gluten-Free") or (MH "Micronutrients") ) OR ( (MH "Dietary Fiber") OR (MH "Cereals+") OR (MH "Dietary Fats+") OR (MH "Calcium, Dietary") OR (MH "Bread") OR (MH "Nutrients+") OR (MH "Nuts+") OR (MH "Seeds+") ) OR ( (MH "Skinfold Thickness") OR (MH "Body Composition+") OR (MH "Metals, Heavy+") OR (MH "Diet+") OR (MH "Malnutrition") OR (MH "Nutrition+") OR (MH "Eating") OR (MH "Probiotics") OR (MH "Prebiotics") OR (MH "Vitamins+") or (MH "Absorptiometry, Photon") ) OR TI ( toxicity or inulin or (Photon Absorptiometry or dexa) or skinfold\* or (body n1 composition) or (arsenic\* or heavy metal\*) or (nutrition\* or diet\* or feed\* or eating or malnutrition\*) or (probiotic\* or prebiotic\* or microbial supplement or vitamin\* or micronutrient or B vitamins or folate or niacin or B12 or riboflavin or calcium or iron or zinc or magnesium or fiber or protein\* or grain\* or seed\* or starch\* or cereal or gliadin\* or glutenin\*) or (nutrition\* or diet\* or feed\* or eating or malnutrition\*) or (nutrition\* or diet\* or feed\* or eating or semolina or durum) ) OR AB ( toxicity or inulin or (Photon Absorptiometry or dexa) or skinfold\* or (body n1 composition) or (arsenic\* or heavy metal\*) or (nutrition\* or titicale or kamut or spelt or semolina or durum) ) OR AB ( toxicity or inulin or (Photon Absorptiometry or dexa) or skinfold\* or (body n1 composition) or (arsenic\* or heavy metal\*) or (nutrition\* or diet\* or feed\* or eating or malnutrition\* or micronutrient or B vitamins or folate or niacin or B12 or riboflavin or calcium or ror or inulin or (Photon Absorptiometry or dexa) or skinfold\* or (body n1 composition) or (arsenic\* or heavy metal\*) or (nutrition\* or diet\* or feed\* or eating or malnutrition\*) or (probiotic\* or prebiotic\* or microbial supplement or vitamin\* or micronutrient or B vitamins or folate or niacin or B12 or riboflavin or calcium or iron or zinc or magnesium or fiber or protein\* or grain\* or seed\* or starch\* or cereal or gliadin\* or gluteni\*) or (gluten\* or oat or Avena or wheat or Triticum o

Hits: 1,859

Embase [August 20, 2018]

1. exp CELIAC DISEASE/

2. (celiac\* or coeliac\*).ti,ab.

3. 1 or 2

4. (nutrition\* or diet\* or feed\* or eating or malnutrition\*).ti,ab.

5. exp DIET/

6. exp nutritional therapy/ or exp diet therapy/

7. exp MALNUTRITION/

8. exp Eating/

9. exp FEEDING BEHAVIOR/

10. exp Gluten/

(continued on next page)

Figure 1. Search strategy for the celiac disease scoping review.

11. exp dietary fiber/
12. (probiotic* or prebiotic* or microbial supplement or vitamin* or micronutrient or B vitamins or folate or niacin or B12 or riboflavin or calcium or iron or zinc or magnesium or fiber or protein* or grain* or seed* or starch* or cereal or gliadin* or glutenin*).ti,ab.
13. exp STARCH/
14. (gluten* or oat or Avena or wheat or Triticum or rye or Secale or barley or Hordeum or triticale or kamut or spelt or semolina or durum).ti,ab.
15. exp plant seed/
16. exp VITAMIN/
17. exp carbohydrate diet/
18. exp trace element/
19. exp probiotic agent/
20. exp MINERAL/
21. exp heavy metal/
22. exp ARSENIC/
23. exp Body Composition/
24. (arsenic* or heavy metal*).ti,ab.
25. (body adj1 composition).ti,ab.
26. bioelectrical impedance analysis.ti,ab.
27. exp impedance/
28. exp SKINFOLD THICKNESS/
29. skinfold*.ti,ab.
30. exp photon absorptiometry/
31. (Photon Absorptiometry or dexa).ti,ab.
32. exp toxicity testing/
33. toxicity.ti,ab.
34. exp INULIN/
35. inulin.ti,ab.
36. or/4-35
37. 3 and 36
38. animals/ not humans/
39. 37 not 38
40. (random\$ or placebo\$ or single blind\$ or double blind\$ or triple blind\$).ti,ab.
41. RETRACTED ARTICLE/
42. (animal\$ not human\$).sh,hw.
43. (book or conference paper or editorial or letter or review).pt. not exp randomized controlled trial/
44. (random sampl\$ or random digit\$ or random effect\$ or random survey or random regression).ti,ab. not exp randomized controlled trial/

(continued on next page)

Figure 1. (continued) Search strategy for the celiac disease scoping review.

45. (40 or 41) not (42 or 43 or 44) 46. exp cohort analysis/ or exp longitudinal study/ or exp prospective study/ or exp follow up/ or cohort\$.tw. or exp case control study/ or (case\$ and control\$).tw. 47. exp review/ or exp meta analysis/ 48. exp Systematic Review/ 49. (literature adj3 review\$).ti,ab. 50. (medline or medlars or embase or pubmed or cinahl or amed or psychit or psychit or psychinfo or psycinfo or scisearch or cochrane).ti,ab. 51. (47 or 48 or 49) and (50 or 41) 52. (systematic\$ adj2 (review\$ or overview)).ti,ab. 53. (meta?anal\$ or meta anal\$ or meta-anal\$ or metaanal\$ or metaanal\$).ti,ab. 54. 39 and (51 or 52 or 53 or 45 or 46) 55. limit 54 to (english language and yr="2007 -Current") Hits: 5465 Medline ovid [July 28, 2018] 1. exp CELIAC DISEASE/ 2. (celiac\* or coeliac\*).ti,ab. 3.1 or 2 4. (nutrition\* or diet\* or feed\* or eating or malnutrition\*).ti,ab. 5. exp DIET/ 6. exp NUTRITION THERAPY/ or exp NUTRITION ASSESSMENT/ 7. exp MALNUTRITION/ 8. exp Eating/ 9. exp FEEDING BEHAVIOR/ 10. exp Glutens/ 11. animals/ not humans/ 12. (probiotic\* or prebiotic\* or microbial supplement or vitamin\* or micronutrient or B vitamins or folate or niacin or B12 or riboflavin or calcium or iron or zinc or magnesium or fiber or protein\* or grain\* or seed\* or starch\* or cereal or gliadin\* or glutenin\*).ti,ab. 13. exp STARCH/ 14. (gluten\* or oat or Avena or wheat or Triticum or rye or Secale or barley or Hordeum or triticale or kamut or spelt or semolina or durum).ti.ab. 15. exp seeds/ 16. exp VITAMINS/ 17. exp dietary carbohydrates/ or exp dietary fiber/ or exp starch/ 18. exp micronutrients/ or vitamins/ 19. exp Probiotics/ 20. exp MINERALS/ (continued on next page)

Figure 1. (continued) Search strategy for the celiac disease scoping review.

21. Metals, Heavy/	]
22. exp ARSENIC/	
23. exp Body Composition/	
24. (arsenic* or heavy metal*).ti,ab.	
25. (body adj1 composition).ti,ab.	
26. bioelectrical impedance analysis.ti,ab.	
27. exp Electric Impedance/	
28. exp SKINFOLD THICKNESS/	
29. skinfold*.ti,ab.	
30. exp Absorptiometry, Photon/	
31. (Photon Absorptiometry or dexa).ti,ab.	
32. exp TOXICITY TESTS/	
33. toxicity.ti,ab.	
34. exp INULIN/	
35. inulin.ti,ab.	
36. or/4-35	
37. 3 and 36	
38. animals/ not humans/	
39. 37 not 38	
40. randomized controlled trial.sh.	
41. controlled clinical trial.pt.	
42. randomized controlled trial*.sh.	
43. random allocation.sh.	
44. double blind method.sh.	
45. single blind method.sh.	
46. clinical trial.pt.	
47. exp Clinical Trial/	
48. (clinical* adj25 trial*).ti,ab.	
49. ((singl* or doubl* or trebl* or tripl*) adj25 (blind* or mask*)).ti,ab.	
50. placebos.sh.	
51. Placebo*.ti,ab.	
52. random*.ti,ab.	
53. research design.sh.	
54. comparative study.sh.	
55. exp evaluation studies/	
56. follow up studies.sh.	
57. prospective studies.sh.	
	continued on next page)
Eigure 1 (continued) Search strategy for the coliac disease scening review	.onanaca on next page)

Figure 1. (continued) Search strategy for the celiac disease scoping review.

58. (control* or prospectiv* or volunteer*).ti,ab.
59. exp cohort studies/
60. cohort*.tw.
61. controlled clinical trial.pt.
62. epidemiological methods/
63. limit 62 to yr=1971-1988
64. exp case-control studies/
65. (case adj2 control*).tw.
66. (review or review,tutorial or review, academic).pt.
67. meta-analysis.pt.
68. meta-analysis.sh.
69. (meta-analys\$ or meta analys\$ or metaanalys\$).tw,sh.
70. (systematic\$ adj5 review\$).tw,sh.
71. "Cross-Sectional Studies"/ or cross sectional.ti,ab.
72. exp Regression Analysis/ or Regression analyses.ti,ab.
73. exp "Surveys and Questionnaires"/
74. (survey* or questionnaire*).ti,ab.
75. exp Regression Analysis/ or (Regression adj1 analyses).ti,ab.
76. or/40-75
77. 39 and 76
78. limit 77 to (english language and yr="2007 -Current")
Hits: 3389
Total hits: 10,810

Figure 1. (continued) Search strategy for the celiac disease scoping review.

examining the validity and reliability of nutrition assessment methods?

- 2. In individuals with CD, what is the availability of the literature examining the effects of different nutrition interventions on nutrition-related health outcomes?
- 3. In individuals with CD, what is the availability of the literature examining the effectiveness of various tools/measures to monitor gluten-free diet adherence/compliance?

Before the start to this scoping review, the authors searched PROSPERO,<sup>9</sup> an international database of prospective systematic reviews in health and other fields, using the terms *celiac* and *coeliac* to identify any potential scoping review with similar scope, but none were identified. There were a few systematic review protocols registered on PROSPERO that may be relevant, and these protocols will be highlighted in the discussion section.

### PROTOCOL

This scoping review used and adapted the methodological framework from the works of Arskey and O'Malley,<sup>10</sup> Levac and colleagues (updated version),<sup>8</sup> and the Joanna Briggs Institute,<sup>11</sup> and also followed the Preferred Reporting Items for Systematic Reviews and Meta-Analyses Extension for Scoping Reviews checklist.<sup>12</sup>

# Search Strategy and Study Selection

An electronic literature search of four databases—Cochrane Database for

systematic reviews, CINAHL (EBSCO), Embase, and Ovid MEDLINE-was conducted in July/August 2018 by a medical librarian using a combination of search terms (Figure 1). The *a priori* eligibility criteria were categorized based on the population, concept, and context mnemonic, as recommended by the Joanna Briggs Institute.<sup>11</sup> The population of this scoping review included any individuals with CD, with no limit on age or sex. The concept related to nutrition care based on the Nutrition Care Process framework<sup>13</sup> (eg, nutrition assessment, intervention, and monitoring). To increase the breadth of this scoping review, the context was left open so evidence could be from any context (eg, setting or geographical locations). Because it is not necessary to specify outcomes for a scoping review,<sup>11</sup> that was left open as well.

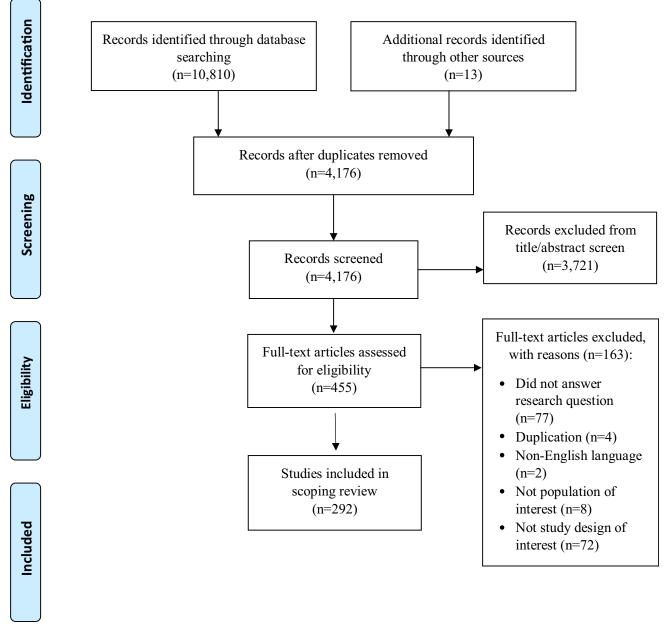


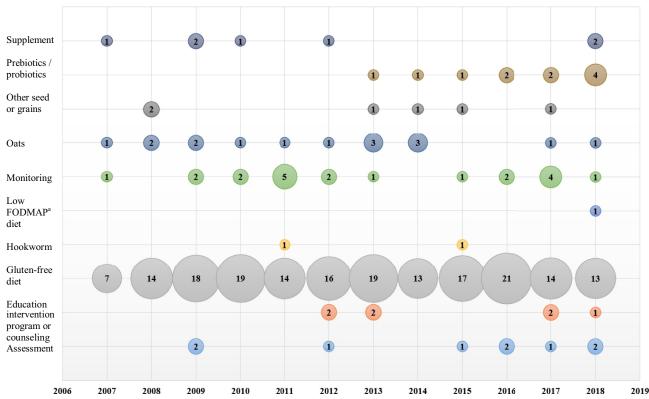
Figure 2. Preferred Reporting Items for Systematic Reviews and Meta-Analyses flow diagram: Celiac disease scoping review.

While a scoping review is broad in nature, it is also important to balance with the availability of resources. Because the goal of this scoping review was to help provide a better direction (eg, whether there is a need/adequate evidence, and/or in which area) when updating the Celiac Disease Guideline<sup>7</sup> by the Academy of Nutrition and Dietetics, the search was limited to any relevant guidelines, consensus statements, recommendations, clinical updates, technical reports, systematic reviews/meta-analysis, experimental trials, and observational studies that were published in English and between 2007 (the literature search for the last CD guideline<sup>7</sup> by the Academy of Nutrition and Dietetics ended in January 2007) and 2018.

For the purpose of this scoping review, guidelines, consensus statements, recommendations, clinical updates, and technical reports published before 2007 were also included in this scoping review to provide better understanding of the breadth and depth of the existing ones that were completed by different organizations. Only the most updated version was included. For example, the American Gastroenterological Association published a technical review on diagnosis and management of CD in 2001 and an update in 2006, only the 2006 version<sup>14</sup> was included.

# Data Extraction and Evidence Mapping

Search results were uploaded to Rayyan,<sup>15</sup> an abstract screening web tool, and screened by EAL staff. After title/abstract and full-text articles screening, information (eg, publication



**Figure 3.** Bubble chart of original research published by year and by topics. The bubble size is proportional to the number of original research studies published in the year and topic. <sup>a</sup>FODMAP=fermentable oligosaccharides, disaccharides, mono-saccharides, and polyols.

year and authors list) from the included articles was exported from Rayyan to Excel (Microsoft). Study design, population (adult, pediatric, or combined), Nutrition Care Process categories (eg, assessment, intervention, or monitoring), subtopics (eg, glutenfree diet, oats, supplements, and prebiotics/probiotics), and outcomes (eg, adherence/management, anthropometrics/growth, bone health, and gastrointestinal symptoms/conditions) were further manually extracted and recorded using the same Excel spreadsheet.

To provide better visualization of the evidence, a bubble chart was used to show the number of original research articles published by year and by topic. For illustrating the distribution of outcomes assessed in the included original intervention studies by study design and by type of intervention, a heat map was used. Lastly, traditional tables were used to show the existing CD guidelines, consensus reports, recommendations, clinical update or report, and technical review and their focuses; relevant systematic review/metaanalysis and outcomes; and original studies examining tools to monitor diet compliance.

#### Consultation

Two content advisors with experience working with CD patients were recruited as volunteers of the Academy of Nutrition and Dietetics to help guide the scoping review process. They were involved with reviewing the initial scoping review search plan to ensure all the necessary search terms were included. They also provided comments on the manuscript.

### **FINDINGS**

The literature search resulted in 10,810 articles with 13 additional studies identified through other sources. A total of 6,647 and 3,721 records were removed because of duplication and lack of relevancy, respectively (Figure 2). Of the 455 full-text articles that were assessed, 292 met the *a priori*  inclusion criteria and were included in this scoping review.

The majority of the studies were observational studies (ie, cohort, casecontrol, cross-sectional, or validation studies) (n=212), followed by experimental trials (ie, randomized or nonrandomized controlled trials or noncontrolled trials) (n=50), guidelines, consensus reports, recommendation statement, clinical update or report, or technical review (n=16), and reviews/meta-analyses systematic (n=14). The number of original research articles by publication year and by topic is illustrated with a bubble chart in Figure 3.

### CD Guidelines, Consensus Reports, Recommendation Statement, Clinical Update or Report, and Technical Review

There were nine CD guidelines, <sup>2-5,7,16-19</sup> three consensus reports, <sup>20-22</sup> one recommendation statement, <sup>23</sup> two clinical updates<sup>24</sup> or report, <sup>25</sup> and one technical review, <sup>14</sup> which were

published between 2004<sup>20</sup> and 2017<sup>2,23</sup> and by various organizations (Table 1). Most of the publications<sup>2,4,5,14,16-21,24,25</sup> covered CD diagnosis and management<sup>7</sup> and 10 of them<sup>2,5,7,14,16-19,24,25</sup> also included monitoring. The three most recent guidelines were published by the National Institute for Health and Care Excellence (2015),<sup>5</sup> Indian Council of Medical Research (2016),<sup>4</sup> and World Gastroenterology Organisation (2017).<sup>2</sup>

### Assessment

Nine observational studies<sup>26-34</sup> focused on assessment; four<sup>26,27,31,34</sup> in adults, three<sup>31-33</sup> in pediatrics, and two<sup>29,30</sup> in both age groups. While six of the included assessment studies<sup>26-29,33,34</sup> aimed to evaluate gluten-free diet adherence, the instruments that were examined varied. For example, some<sup>26,27</sup> developed and validated a gluten-free diet score, while others<sup>28,29,33,34</sup> focused on a new or an adopted food frequency questionnaires or a simple questionnaire/survey. The objectives of the other three studies were slightly different: one<sup>30</sup> developed and validated a scale to evaluate specific self-efficacy (determinants in gluten-free diet adherence), another article<sup>32</sup> focused on the development and validation of the Celiac Disease-Children's Activities Report for advocating self-management. The last study<sup>31</sup> evaluated a modified version of the Italian European Prospective Investigation into Cancer and Nutrition Food Frequency Questionnaire to assess the overall nutrient intake among individuals with CD.

### Intervention

Included studies investigated various nutrition interventions in patients with CD, such as education intervention program or counseling; low fermentable oligosaccharides, disaccharides, monosaccharides, and polyols (FOD-MAP) diet; gluten-free diet; and supplements. The distribution of outcomes assessed in the original intervention studies by study design and by type of intervention is illustrated with a heat map (Figure 4).

**Education Intervention Program or Counseling.** Five experimental trials<sup>35-39</sup> and two observational studies<sup>40,41</sup> investigated the effects of education intervention program or counseling on various health outcomes, but the types of intervention and the components that they entailed were heterogeneous in nature. One study<sup>40</sup> conducted a cross-sectional survey to assess the effects of a single educational meeting on disease knowledge and awareness among adults with CD. The other observational study<sup>41</sup> also used a survey to examine whether dietitian use (ie, the number of times the participant had seen a dietitian) had positive effects on quality of life, severity of symptom, and adherence in adult CD patients.

Different types of interventions were noted in the five included experimental trials for individuals with CD<sup>35-39</sup>: inperson educational program for adults,<sup>36,37</sup> online educational program for adults,<sup>38</sup> text message intervention for both children and adults,<sup>35</sup> and online follow-up consultations for both children and adults.<sup>39</sup> These studies assessed various outcomes, including adherence/management,<sup>35,38</sup> gastrointestinal symptoms and conditions,<sup>35,36</sup> immunology/serology measures,<sup>35,39</sup> mental/cognitive health,<sup>35,37</sup> and quality of life (Figure 4).<sup>35,37,39</sup>

Gluten-Free Diet. Eight systematic review/meta-analysis,<sup>42-49</sup> 9 experimental trials,<sup>50-58</sup> and 176 observational studies<sup>59-234</sup> examined the effects of gluten-free diet. Most of the systematic reviews/meta-analyses<sup>44-49</sup> were published between 2015<sup>44,45</sup> and 2018.48,49 They covered various outcomes, such as bone health,<sup>44</sup> mental health,<sup>45,49</sup> gastrointestinal symptoms/conditions,47 heart health,<sup>48</sup> and quality of life (Table 2).<sup>46</sup> The included experimental trials were published between 2007<sup>50</sup> and 2016,<sup>56,57</sup> with a majority of the studies focusing on adults with CD.<sup>50-53,55,57,58</sup> They assessed a range of outcomes, including anthropometrics/ growth,<sup>51,52,54</sup> bone health,<sup>51,52,58</sup> gastrointestinal symptoms/conditions, 50-52,57 immunology/serology measures,<sup>50,52</sup> mental/cognitive health, 51-53 neuro-/ autoimmune diseases,55 nutritional status,<sup>51,52,56</sup> and quality of life (Figure 4).<sup>51</sup> Of the 176 observational studies, 44% focused on adults, 44% on children, and the remaining 12% included both age groups. All of the outcomes listed in Figure 4, except for nutrition knowledge/ awareness, were assessed by the included observational studies.

**Hookworm.** Two experimental trials<sup>235,236</sup> examined whether the use of hookworm infection could be an effective treatment strategy for CD patients. Both studies recruited adult patients, with the length of study ranging from 21<sup>236</sup> to 52<sup>235</sup> weeks. After inoculation, participants underwent gluten/ wheat challenge in both studies. Similar outcomes, such as gastrointestinal symptoms and conditions, immunology/serology measures, and quality of life, were collected (Figure 4).

**Low FODMAP Diet.** Only one intervention study<sup>237</sup> examined the effectiveness of a low FODMAP diet in CD patients who are on a gluten-free diet. These adult participants were educated on a low FODMAP diet in the beginning and were asked to follow this diet throughout the study. Outcomes, such as quality of life, mental health, and gastrointestinal symptoms, were assessed at baseline, 1 month, and 3 months (Figure 4).

Oats. Of the 21 included studies, 5<sup>238-242</sup> were systematic reviews/metaanalysis, 8<sup>243-250</sup> were experimental trials, and 8<sup>205,251-257</sup> were observational studies. Three of the systematic reviews/meta-analysis were published in 2016<sup>238,240</sup> or 2017.<sup>241</sup> They included both children and adults with dermatitis herpetiformis, gastrointestinal symptoms/conditions, and/or immunology/serology measures as the outcomes of interest (Table 2). The included experimental studies were published between 2008<sup>246</sup> and 2018<sup>248</sup> with two studies<sup>245,246</sup> from the same trial. Four<sup>244,247,248,250</sup> of the studies focused on children; two<sup>243,249</sup> focused on adults and two<sup>245,246</sup> included both age groups. Outcomes, such as gastrointestinal symptoms/ conditions<sup>244,246-249</sup> and immunology/ serology measures,<sup>243,244,246-248</sup> were reported by most of those trials (Figure 4). Most of the observational studies<sup>205,252-255</sup> focused on the adult population; two<sup>256,257</sup> included children and one<sup>251</sup> included both age groups. Similar to experimental trials, most observational studies collected outcomes on gastrointestinal symptoms/conditions<sup>205,251,253,256,257</sup> and immunology/serology measures<sup>251,252,254</sup> (Figure 4).

**Other Seed or Grains.** Six studies examined whether consuming other

Year	Association (reference)	Title	Туре	Focus
2004	National Institutes of Health (NIH) <sup>20</sup>	NIH Consensus Development Conference on Celiac Disease	Consensus report	Celiac disease diagnosis; management
2005	North American Society for Pediatric Gastroenterology, Hepatology and Nutrition <sup>16</sup>	Guideline for the diagnosis and treatment of celiac disease in children: Recommendations of the North American Society for Pediatric Gastroenterology, Hepatology and Nutrition	Guideline	Celiac disease diagnosis; management; monitoring
2006	American Gastroenterological Association (AGA) <sup>14</sup>	American Gastroenterological Association (AGA) Institute technical review on the diagnosis and management of celiac disease	Technical review	Celiac disease diagnosis; management; monitoring
2007	Gastroenterological Society of Australia <sup>24</sup>	<i>Coeliac Disease</i> , 4 <sup>th</sup> edition	Clinical update	Celiac disease diagnosis; management; monitoring
2008	Federation of International Societies of Pediatric Gastroenterology, Hepatology, and Nutrition <sup>21</sup>	Federation of International Societies of Pediatric Gastroenterology, Hepatology, and Nutrition consensus report on celiac disease	Consensus report	Celiac disease diagnosis; management
2009	Academy of Nutrition and Dietetics <sup>7</sup>	Celiac Disease Guideline	Guideline	Nutrition assessment; management; monitoring
2012	European Society for Pediatric Gastroenterology, Hepatology, and Nutrition guidelines <sup>3</sup>	European Society for Pediatric Gastroenterology, Hepatology, and Nutrition guidelines for the diagnosis of coeliac disease	Guideline	Celiac disease diagnosis
2013	American College of Gastroenterology (ACG) <sup>17</sup>	ACG clinical guidelines: diagnosis and management of celiac disease	Guideline	Celiac disease diagnosis; management; monitoring
2013	British Society of Paediatric Gastroenterology, Hepatology and Nutrition (BSPGHAN); Coeliac UK <sup>18</sup>	Joint BSPGHAN and Coeliac UK guidelines for the diagnosis and management of coeliac disease in children	Guideline	Celiac disease diagnosis; management; monitoring
2014	British Society of Gastroenterology <sup>19</sup>	Diagnosis and management of adult coeliac disease: guidelines from the British Society of Gastroenterology	Guideline	Celiac disease diagnosis; management; monitoring
2015	National Institute for Health and Care Excellence <sup>5</sup>	Coeliac Disease: Recognition, Assessment and Management	Guideline	Celiac disease diagnosis; management; monitoring
2016	Association of European Coeliac Societies; US Celiac Disease Foundation <sup>22</sup>	Transition from childhood to adulthood in coeliac disease: the Prague consensus report	Consensus report	Management (transition from childhood to adulthood in celiac disease)
2016	Indian Council of Medical Research (ICMR) <sup>4</sup>	ICMR Guideline on Diagnosis and Management of Celiac Disease	Guideline	Celiac disease diagnosis; management (continued on next page)

Year	Association (reference)	Title	Туре	Focus
2016	North American Society for Pediatric Gastroenterology, Hepatology and Nutrition (NASPGHAN) <sup>25</sup>	NASPGHAN Clinical Report on the Diagnosis and Treatment of Gluten-related Disorders	Clinical report	Celiac disease diagnosis; management; monitoring
2017	US Preventive Services Task Force <sup>23</sup>	Screening for Celiac Disease: US Preventive Services Task Force Recommendation Statement	Recommendation statement	Celiac disease screening
2017	World Gastroenterology Organisation <sup>2</sup>	World Gastroenterology Organisation Global Guidelines: Celiac Disease February 2017	Guideline	Celiac disease diagnosis; management; monitoring

Table 1. Celiac disease guidelines, consensus reports, recommendation, clinical update or report, and technical review (continued)

		ation inter ram or cou		Gh	iten-free	diet		Hookwor	m	Low	FODMA	P diet		Oats		Othe	r seed or g	grains	Prebio	otics / pro	biotics	s	upplemen	ıt
	ET	OS	Т	ET	OS	Т	ET	OS	Т	ET	OS	Т	ET	OS	Т	ET	OS	Т	ET	OS	Т	ET	OS	Т
Total number of studies by ET, OS, T	5	2	7	9	176	185	2	0	2	1	0	1	8	8	16	5	1	6	11	0	11	7	0	7
Adherence / management	2	1	3	0	1	1	0	0	0	0	0	0	0	1	1	0	0	0	0	0	0	0	0	0
Anthropometrics / growth	0	0	0	3	30	33	0	0	0	0	0	0	1	0	1	0	1	1	0	0	0	0	0	0
Bone health	0	0	0	3	18	21	0	0	0	0	0	0	0	1	1	0	0	0	0	0	0	1	0	1
Cancer	0	0	0	0	3	3	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dermatitis herpetiformis	0	0	0	0	3	3	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Endocrine (e.g., HbA1c, glucose, etc.)	0	0	0	0	16	16	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Eye health	0	0	0	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Gastrointestinal bacterial / gut microbiota	0	0	0	0	4	4	0	0	0	0	0	0	1	0	1	0	0	0	7	0	7	0	0	0
Gastrointestinal symptoms / conditions	2	1	3	4	45	49	2	0	2	1	0	1	5	5	10	4	1	5	3	0	3	1	0	1
Heart health (e.g., lipid panel, homocysteine levels, etc.)	0	0	0	0	15	15	0	0	0	0	0	0	0	0	0	1	0	1	0	0	0	2	0	2
Immunology / serology measures	2	0	2	2	21	23	2	0	2	0	0	0	5	3	8	2	0	2	5	0	5	0	0	0
Liver health (e.g., liver panel / disease)	0	0	0	0	4	4	0	0	0	0	0	0	0	0	0	0	1	1	0	0	0	0	0	0
Mental / cognitive health	2	0	2	3	32	35	0	0	0	1	0	1	0	2	2	0	0	0	0	0	0	1	0	1
Metal / mycotoxin accumulation	0	0	0	0	2	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Mortality	0	0	0	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Neuro / autoimmune diseases	0	0	0	1	8	9	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Nutrition knowledge / awareness	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Nutritional quality / diet quality	0	0	0	0	14	14	0	0	0	0	0	0	1	1	2	0	0	0	0	0	0	0	0	0
Nutritional status	0	0	0	3	28	31	0	0	0	0	0	0	0	1	1	0	0	0	0	0	0	3	0	3
Oral health	0	0	0	0	2	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Quality of life (e.g., perceived / functional health, etc.)	3	1	4	1	36	37	1	0	1	1	0	1	0	1	1	0	0	0	1	0	1	1	0	1
Reproductive health	0	0	0	0	4	4	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Sleep outcome	0	0	0	0	3	3	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Urinary health	0	0	0	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

Figure 4. The distribution of outcomes assessed in the included original intervention studies by study design and type of intervention is illustrated with a heat map. Red=highest number of studies. Yellow=number of studies at around the 50th percentile. Green=lowest number of studies. ET=experimental trial. OS=observational studies. T=total number of studies within each type of intervention.

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seed or grains—quinoa,<sup>258</sup> teff,<sup>259</sup> *Triticum monococcum*,<sup>260,261</sup> and wheatbased starch hydrolysates<sup>262,263</sup> could negatively affect health outcomes among CD patients. Except for one observational study,<sup>259</sup> all of them<sup>258,260-263</sup> were experimental trials. Three studies<sup>258,259,262</sup> focused on the adult populations, while the other three studies<sup>260,261,263</sup> included both children and adults. Outcomes assessed are displayed in Figure 4.

Probiotics/Prebiotics. Of 11 the experimental trials included, 2 studies<sup>264,265</sup> (from the same trial) effects investigated the of oligofructose-enriched inulin, and the other 9 studies<sup>266-274</sup> investigated the effect of either a single or a mixture of different probiotic studies<sup>266,267,269,273,274</sup> strains. Five included adult patients and the rest of the studies<sup>264,265,268,270-272</sup> focused on the pediatric population. Various outcomes, such as gastrointestinal bacterial/gut microbiota, gastrointestinal symptoms/conditions, immunology/ serology measures, and quality of life, were collected (Figure 4).

**Supplements.** Different types of supplements were studied in the 7 included experimental trials: L-carnitine,<sup>275</sup> iron,<sup>276</sup> B vitamins,<sup>277,278</sup> zinc,<sup>279,280</sup> and calcium and alfacalcidol.<sup>58</sup> Five<sup>58,275-278</sup> of the studies focused on adults, while the other two<sup>279,280</sup> studied the pediatric population. These studies collected different outcomes, such as bone health,<sup>280</sup> gastrointestinal symptoms and conditions,<sup>276</sup> plasma homocysteine,<sup>277,278</sup> mental/ cognitive health,<sup>275</sup> nutritional status (eg, hemoglobin, iron, ferritin, zinc, and copper levels),<sup>276,279,280</sup> and quality of life (Figure 4).<sup>275</sup>

Monitoring. One systematic review/ meta-analysis<sup>281</sup> examined the sensitivity and specificity of serum transglutaminase endomysial and antibodies on detecting patients with villous atrophy (Table 2). There were 21 original studies<sup>33,141,282-300</sup> investigating the effectiveness of different tools/measures, such as immunoglobulin G anti-tissue deamidated gliadin peptides and immunoglobulin A antitissue transglutaminase, on monitoring adherence/compliance. Nine studies<sup>282-284,286,289,291,293,295,297</sup> focused on adults, six studies<sup>33,141,287,288,294,298</sup> focused on children, and six studies<sup>285,290,292,296,299,300</sup> included on both age groups (Table 3).

### Summary

The goal of this scoping review was to gain a better understanding of the landscape of CD literature, which could help to inform the need/scope of the development of future systematic reviews and evidence-based practice guidelines in this area.

One of the reasons to include guidelines as part of the search was to examine whether there are recently published guidelines with similar scope and methodological rigor that could be adopted/recommended. In fact, this scoping review found several recent CD guidelines that were developed by different organizations, that is, the National Institute for Health and Care Excellence (2015),<sup>5</sup> Indian Council of Medical Research (2016),<sup>4</sup> and the World Gastroenterology Organisation (2017).<sup>2</sup> All three guidelines included some aspects of nutrition care and the role of dietitians in this process (eg. "Lifelong and complete avoidance of gluten or gluten containing dietary items is the most effective and the main stay of treatment of CeD. While planning gluten-free diet, all patients should be counselled for a balanced diet as per their nutritional requirement."<sup>4</sup>). However, different methods in terms of literature search and assessment of evidence strength were used. This underscores the importance of careful consideration and using a tool, such as AGREE II (Appraisal of Guidelines Research and Evaluation II).<sup>301</sup> to evaluate the methodological quality of existing guidelines even when they seem to be appropriate and applicable to registered dietitian nutritionists when deciding whether they could be adopted or recommended.

The search results for assessment studies presented an interesting phenomenon. Most of the included studies investigated different tools/ways to assess gluten-free diet adherence, yet none of them directly examined the validity and reliability of an assessment tool for nutrition status. A similar situation also was noted in the last Celiac Disease Guideline<sup>7</sup> by the Academy of Nutrition and Dietetics. For example, there was a recommendation for "assessing biochemical data and results of medical procedures: the registered dietitian (RD) should assess the biochemical data and review the results of medical procedures in individuals with celiac disease. regardless of presentation and clinical symptoms, including (but not limited to) the following: Gastrointestinal profile [e.g., intestinal biopsy (or skin biopsy in the case of dermatitis herpetiformis) and celiac antibodies]; Nutritional anemia profile (e.g., folate, ferritin and vitamin B12); Vitamin profile (e.g., thiamin, vitamin B6 and 25-hydroxy vitamin D); Mineral profile (e.g., copper and zinc); Lipid profile; Electrolyte and renal profile." The supporting evidence was from intervention studies that had examined the effects of gluten-free diet among individuals with CD.<sup>7</sup> It is also important to note that most of the adherence studies focused on identifving different factors that are associated with adherence, which is not the goal of this scoping review, instead of an actual adherence assessment tool that practitioners can use. Therefore, those studies were excluded.

There were a total of 235 original intervention studies that met the eligibility criteria. Of those included, the majority ( $\sim$ 79%) examined the effects of gluten-free diet, followed by  $(\sim 7\%)$ , prebiotics/probiotics oats  $(\sim 4\%)$ , education intervention program or counseling ( $\sim$  3%), supplement  $(\sim 3\%)$ , other seed or grains  $(\sim 3\%)$ , hookworm (<1%), and low FODMAP diet (<1%). The fact that most of the studies focused on gluten-free diet aligns with the unequivocal recommendation that lifelong glutenfree diet is the most effective treatment for CD.<sup>4,5</sup> Thus, it appears that most of the research in this area focused on the effect of gluten-free diet on various health outcomes and nutrition status (eg. bone health), with a secondary goal to explore whether individuals with CD would require additional assessment (eg, iron status) or intervention as a result of this restrictive diet. Similarly, when searching in PROSPERO for similar reviews in the beginning of the project, most of the potentially relevant systematic review protocols were related to gluten-free diet. In fact, three were identified: two<sup>302,303</sup> focused on the effect of gluten-free diet on bone health and

one<sup>304</sup> focused on its effect on body mass.

The second category with the most studies was oats. According to the National Institute for Health and Care Excellence, individuals "can choose to include gluten-free oats in their diet at any stage and they will be advised whether to continue eating gluten-free oats depending on their immunological, clinical or histological response." Similarly, the World Gastroenterology Global Organisation Guidelines mentioned that "oats may he consumed," but cautioned about potential contamination with wheat.<sup>2</sup> The Indian Council of Medical Research,<sup>4</sup> on the other hand, recommended to completely avoid oats, as they are often contaminated with wheat. The differences in recommendations seem to be due to the availability of gluten-free oats in those geographical locations instead of based on the effect of consumption of gluten-free oats. In fact, three of the most recent systematic reviews/meta-analysis<sup>238,240,241</sup> supported this rationale and found that the consumption of gluten-free oats was generally safe but some oat cultivars may produce an immune reaction in those who are more sensitive.<sup>238</sup> Thus, as recommended by the National Institute for Health and Care Excellence, long-term regular followup is the key.

Prebiotics/probiotics was next on the list; no systematic review/metaanalysis was found in the search and this topic was also not mentioned in the three recent guidelines.<sup>2,4,5</sup> This may be due to the lack of literature during the development of those guidelines, considering the first study on this topic was published in 2013, given our search criteria included articles published betwen January 2007 and August 2018.<sup>273</sup> However, there was a systematic review protocol<sup>305</sup> on the clinical efficacy and safety of probiotics among children with gastrointestinal conditions, including CD, registered on PROSPERO in 2016. It appears that this review has not been started and no published article was found.

Education intervention program or counseling is important to help improve diet adherence when glutenfree diet is the most effective treatment for CD. However, it is important to note that the included studies focused on different types of intervention program (eg, online, text message, and in person), potentially increasing the difficulty of synthesizing those data. The search in PROSPERO identified two prospective systematic reviews that may fall under this category, both systematic review protocols were registered in 2018 and are still in progress. One<sup>306</sup> is evaluating the effectiveness of mobile and web-based applications support to selfmanagement during transition from pediatric to adult among those with chronic illness, including CD. The other systematic review<sup>307</sup> is focusing on determining which behavior-change techniques could improve adherence in patients with CD.

A similar situation was also noted with supplement and other seed or grains. Although approximately 3% of the studies examined each of these two topics, the actual intervention differed. For instance, of the seven experimental trials included in the supplement category, only one study looked at each of these supplements: L-carnitine, iron, and calcium+alfacalcidol: and two studies looked each of these supplements: B vitamins and zinc. The last two topics (ie, hookworm and low FODMAP diet) are interesting types of intervention for CD, yet the number of the included studies was sparse.

The third research question was about examining tools/measures to monitor adherence/compliance among individuals with CD. Of the three most recent guideline, the World Gastroenterology Organisation Global Guidelines<sup>2</sup> mentioned that "studies suggest that periodic testing for IgA anti-tTG or IgA anti-DGP is the preferred method for monitoring compliance," the Indian Council of Medical Research<sup>4</sup> recommended "serological tests at 6 months and one year can be used to monitor adherence," but did not specify one. Lastly, the National Institute for Health and Care Excellence<sup>5</sup> recommended not using "serological testing alone to determine whether gluten has been excluded from the person's diet" because low- or very-low-quality evidence was available to support the use of various serology measures to monitor compliance. Similar to the results found in this scoping review, although 21 original studies focused on

this area, they differed in their tools/ measures of interest, as well as in their methods to assess those tools/measures (eg, comparison with reference standard and control). For example, of the 21 studies, only 1 study<sup>292</sup> examined the effectiveness of using gliadin 33-mer-equivalent peptidic epitopes in human feces to monitor compliance. The heterogeneity and availability of studies for each of the serology measures may potentially result in low- or very-low-quality evidence, as seen in the CD guideline published by the National Institute for Health and Care Excellence.<sup>5</sup>

### Strengths and Limitations

There are several strengths in this scoping review: content advisors reviewed the initial scoping review search plan to ensure that all of the relevant search terms were included and a medical librarian conducted a comprehensive literature search in four databases and tailored search strategy/ term specific to each database, EAL staff used and adapted a methodological framework based on the works of Arskey and O'Malley,<sup>10</sup> Levac and colleagues (updated version),8 and the Joanna Briggs Institute,<sup>11</sup> and this scoping review also followed the Preferred Reporting Items for Systematic Reviews and Meta-Analyses Extension for Scoping Reviews checklist.

Several limitations should be noted. While it is preferred to have two reviewers screen through all of the abstracts from the beginning, only one reviewer screened the abstracts during the title/abstract screening round due to the lack of resources (eg, financial). However, to address this limitation, extreme cautious was exercised to ensure that only the "true" excludes were being eliminated during this first round. Any unsure ones were treated as "includes" and went to the full-text screening round, where two reviewers reviewed all of the included studies. Although the search plan was comprehensive, there is always a possibility that it did not capture all of the studies that met the inclusion criteria. However, efforts were made to look for potential studies through the existing included ones and content advisors also reviewed the manuscript.

### Table 2. Relevant celiac disease systematic review or meta-analysis published between 2007 and 2018

Intervention	Author		
and year	(reference)	Title	Outcomes
Gluten-free diet			
2008	Akobeng <sup>42</sup>	Systematic review: Tolerable amount of gluten for people with coeliac disease	Threshold of tolerable amount of gluten
2008	Haines <sup>43</sup>	Systematic review: The evidence base for long-term management of coeliac disease	Management of celiac disease (eg, the effect of gluten-free diet on various health outcomes)
2015	Grace-Farfaglia <sup>44</sup>	Bones of contention: Bone mineral density recovery in celiac disease—A systematic review	Bone mineral density
2015	Zingone <sup>45</sup>	Psychological morbidity of celiac disease: A review of the literature	Anxiety, depression, fatigue
2017	Burger <sup>46</sup>	Systematic review with meta-analysis: Dietary adherence influences normalization of health-related quality of life in coeliac disease	Health-related quality of life
2017	Szakacs <sup>47</sup>	Younger age at diagnosis predisposes to mucosal recovery in celiac disease on a gluten-free diet: A meta-analysis	Mucosal recovery
2018	Potter <sup>48</sup>	Effect of the gluten-free diet on cardiovascular risk factors in patients with coeliac disease: A systematic review	Cardiovascular risk factors
2018	Sainsbury <sup>49</sup>	The relationship between gluten free diet adherence and depressive symptoms in adults with coeliac disease: A systematic review with meta-analysis	Depressive symptoms
Monitoring			
2017	Silvester <sup>281</sup>	Tests for serum transglutaminase and endomysial antibodies do not detect most patients with celiac disease and persistent villous atrophy on gluten-free diets: A meta-analysis	Sensitivity and specificity of serum transglutaminase and endomysial antibodies in detecting individuals with villous atrophy
Oats			
2007	Garsed <sup>239</sup>	Can oats be taken in a gluten-free diet? A systematic review	Gastrointestinal symptoms/conditions, immunology/ serology measures
2009	Pulido <sup>242</sup>	Introduction of oats in the diet of individuals with celiac disease: A systematic review	Gastrointestinal symptoms/conditions, immunology/ serology measures
2016	de Souza <sup>238</sup>	Pure oats as part of the Canadian gluten-free diet in celiac disease: The need to revisit the issue	Gastrointestinal symptoms/conditions, immunology/ serology measures
2016	La Vieille <sup>240</sup>	Celiac disease and gluten-free oats: A Canadian position based on a literature review	Gastrointestinal symptoms/conditions, immunology/ serology measures
2017	Pinto-Sanchez <sup>241</sup>	Safety of adding oats to a gluten-free diet for patients with celiac disease: Systematic review and meta-analysis of clinical and observational studies	Dermatitis herpetiformis, gastrointestinal symptoms/ conditions, immunology/serology measures

Table 3. Original research studies examining tools to monitor diet compliance

Year	Author (reference)	Title	Methods to assess the tools/measures (eg, comparison with reference standard, control)	Tools/measures
2007	Leffler <sup>282</sup>	A prospective comparative study of five measures of gluten-free diet adherence in adults with coeliac disease	Expert nutritionist evaluation	Self-reported adherence IgG <sup>a</sup> anti-DGP <sup>b</sup> IgA <sup>c</sup> anti-DGP IgG-IgA anti-DGP IgA anti-t <sup>d</sup> TG <sup>e</sup>
2009	da Silva Kotze <sup>283</sup>	A Brazilian experience of the self transglutaminase-based test for celiac disease case finding and diet monitoring	Upper gastrointestinal endoscopy with duodenal biopsies and IgA EmA <sup>f</sup>	lgA anti-tTG (rapid)
2009	Leffler <sup>284</sup>	A validated disease-specific symptom index for adults with celiac disease	Test-retest reliability; second validation study (SF-36 <sup>9</sup> and EQ-5D <sup>h</sup> )	CSI <sup>i</sup>
2010	Koskinen <sup>285</sup>	Usefulness of small-bowel mucosal transglutaminase-2 specific autoantibody deposits in the diagnosis and follow-up of celiac disease	Conventional celiac serology; mucosal morphology and to the density of IELs <sup>j</sup>	lgA-anti-TG2 <sup>k</sup>
2010	Sugai <sup>286</sup>	Dynamics of celiac disease-specific serology after initiation of a gluten-free diet and use in the assessment of compliance with treatment	Compliance assessment based on opinion of physician, dietitians, and self-reported	IgA anti-tTG IgA anti-DGP IgG anti-DGP IgG-IgA anti-DGP DPG/tTG Screen IgA AGA <sup>I</sup> IgA AAA <sup>m</sup> IgA EmA
2011	Laadhar <sup>287</sup>	Is the rapid whole blood test useful for diagnosis and monitoring celiac disease in children?	IgA anti-tTG (ELISA <sup>n</sup> )	lgA anti-tTG (rapid)
2011	Monzani <sup>288</sup>	Use of deamidated gliadin peptide antibodies to monitor diet compliance in childhood celiac disease	Compliance assessment by investigator or calculated cutoff based on ROC <sup>o</sup> curve	lgA anti-DGP lgG-lgA anti-DGP lgA anti-tTG lgA AGA
2011	Nachman <sup>289</sup>	Serological tests for celiac disease as indicators of long-term compliance with the gluten-free diet	Compliance assessment based on clinical assessment and self-reported; calculated cutoff based on ROC curve and manufacturer	IgA anti-tTG IgA anti-DGP IgG anti-DGP IgG-IgA anti-DGP DPG/tTG Screen IgA AAA IgA EmA (continued on next page)

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Table 3. Original research studies examining tools to monitor diet compliance (contin	ued)
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Year	Author (reference)	Title	Methods to assess the tools/measures (eg, comparison with reference standard, control)	Tools/measures
2011	Planas <sup>290</sup>	Regenerating gene lalpha is a biomarker for diagnosis and monitoring of celiac disease: A preliminary study	lgA anti-tTG and IgA AGA	REG1 $\alpha^{p}$
2011	Purnak <sup>291</sup>	Mean platelet volume could be a promising biomarker to monitor dietary compliance in celiac disease	Control group, level of adherence	MPV <sup>q</sup>
2012	Comino <sup>292</sup>	Monitoring of gluten-free diet compliance in celiac patients by assessment of gliadin 33- mer equivalent epitopes in feces	Different levels to gluten ingestion	33EPs <sup>r</sup>
2012	Leffler <sup>293</sup>	Open conformation tissue transglutaminase testing for celiac dietary assessment	Adherence determined by dietitian	O-tTG <sup>s</sup> C-tTG <sup>t</sup>
2013	Aita <sup>294</sup>	Chemiluminescence and ELISA-based serum assays for diagnosing and monitoring celiac disease in children: A comparative study	ELISAs: lgA anti-tTG lgG anti-tTG lgA anti-DGP lgG anti-DGP lgG-lgA anti-DGP	Chemiluminescent assays: IgA anti-tTG IgG anti-tTG IgA anti-DGP IgG anti-DGP IgG-IgA anti-DGP
2015	Srinivasan <sup>295</sup>	Usefulness of recombinant gamma-gliadin 1 for identifying patients with celiac disease and monitoring adherence to a gluten-free diet	lgA anti-tTG2 lgA anti-DGP lgG anti-DGP	GG1 <sup>u</sup>
2016	Comino <sup>296</sup>	Fecal gluten peptides reveal limitations of serological tests and food questionnaires for monitoring gluten-free diet in celiac disease patients	Dietary compliance; IgA anti-tTG; IgA anti-DGP	GIP <sup>∨</sup>
2016	Lind <sup>297</sup>	Plasma alkylresorcinols reflect gluten intake and distinguish between gluten-rich and gluten- poor diets in a population at risk of metabolic syndrome	Gluten-rich and gluten-poor diet	Plasma total alkylresorcinol concentrations
2017	Adriaanse <sup>298</sup>	Progress towards non-invasive diagnosis and follow-up of celiac disease in children; a prospective multicentre study to the usefulness of plasma I-FABP <sup>W</sup>	Gluten-free diet (26 wk), histological and serological disease markers.	Plasma I-FABP
				(continued on next page)

Table 3. Original research studies examining tools to monitor diet compliance (continued)

Year	Author (reference)	Title	Methods to assess the tools/measures (eg, comparison with reference standard, control)	Tools/measures
2017	Lau <sup>299</sup>	The role of an IgA/IgG-deamidated gliadin peptide point-of-care test in predicting persistent villous atrophy in patients with celiac disease on a gluten-free diet	Duodenal histology	lgA anti-tTG IgA EmA IgG-IgA anti-DGP (Simtomax)
2017	Leonard <sup>141</sup>	Value of IgA tTG in predicting mucosal recovery in children with celiac disease on a gluten-free diet	Duodenal histology	lgA anti-tTG
2017	Moreno <sup>300</sup>	Detection of gluten immunogenic peptides in the urine of patients with coeliac disease reveals transgressions in the gluten-free diet and incomplete mucosal healing	Gluten ingestion (25 and 50 mg)	GIP
2018	Wessels <sup>33</sup>	Assessment of dietary compliance in celiac children using a standardized dietary interview	Standardized dietary interview	lgA-anti-TG2

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<sup>a</sup>lgG=immunoglobulin G. <sup>b</sup>DGP=deamidated gliadin peptide. <sup>c</sup>lgA=immunoglobulin A. <sup>d</sup>anti-t=anti-tissue. <sup>e</sup>TG=transglutaminase. <sup>f</sup>lgA EmA=antiendomysial antibody. <sup>9</sup>SF-36=36-Item Short Form Health Survey. <sup>h</sup>EQ-5D=EuroQol-5 Dimension. <sup>i</sup>CSI=Celiac Symptom Index (disease-specific). <sup>j</sup>IEL=intraepithelial lymphocyte. <sup>k</sup>TG2=transglutaminase-2. <sup>I</sup>IgA AGA=IgA type antigliadin antibody. <sup>m</sup>IgA AAA=IgA isotype antiactin antibody. <sup>n</sup>ELISA=enzyme-linked immunosorbent assay. <sup>o</sup>ROC=receiver operating characteristic curve. <sup>P</sup>REG1 $\alpha$ =regenerating gene 1 $\alpha$ . <sup>q</sup>MPV=mean platelet volume. <sup>r</sup>33Eps=gliadin 33-mer equivalent peptidic epitopes in human feces. <sup>s</sup>O-tTG=stabilized open (active) conformation tTG. <sup>t</sup>C-tTG=closed or undefined conformation tTG. <sup>u</sup>GG1= $\gamma$ -gliadin 1. <sup>v</sup>GIP=gluten immunogenic peptide. <sup>w</sup>I-FABP=intestinal-fatty acid binding protein.

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# CONCLUSIONS AND FUTURE DIRECTION

This scoping review completed a comprehensive literature search to examine the availability of literature on the validity and reliability of nutrition assessment methods, nutrition interventions, and tools/measures to monitor adherence/compliance among individuals with CD.

Based on the scoping review, some topics (eg, gluten-free diet, oats) were already covered by either a recent guideline or systematic review. For instance, a 2017 systematic review on consumption of  $oat^{241}$  included 12 of the 16 original studies founded in this scoping review. Thus, the expert panel on a future CD systematic review/ guideline project can evaluate whether to use this type of recent systematic review by examining the scope and the rigor of the methodology used. A similar approach can be implemented when determining whether a recent guideline can be adapted or recommended. Other topics (eg. prebiotics/ probiotics, education program/counseling focused interventions) that have not been covered by a recent review or guideline could be potential areas to focus on in a new CD systematic review/guideline project. However, consideration also should be balanced with the number of available studies in certain topics, such as with low FOD-MAP diet.

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#### **AUTHOR INFORMATION**

F. W. Cheng is a nutrition researcher and D. Handu is senior scientific director, Evidence Analysis Center, Academy of Nutrition and Dietetic, Chicago, IL.

Address correspondence to: Feon W. Cheng, PhD, MPH, RDN, CHTS-CP, Evidence Analysis Center, Academy of Nutrition and Dietetics, 120 S Riverside Plaza, Suite 2190, Chicago, IL 60606-6995. E-mail: <a href="https://doi.org/10.1071/journal.pdf">https://doi.org/10.1071/journal.pdf</a> (https://doi.org/10.1071/journal.pdf) (https://doi.org/

#### STATEMENT OF POTENTIAL CONFLICT OF INTEREST

No potential conflict of interest was reported by the authors.

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