Cholesterol-lowering effects of oat β-glucan: a meta-analysis of randomized controlled trials

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ABSTRACT

Background: Health claims regarding the cholesterol-lowering effect of soluble fiber from oat products, approved by food standards agencies worldwide, are based on a diet containing ≥3 g/d of oat β-glucan (OBG). Given the number of recently published randomized controlled trials (RCTs), it is important to update the findings of previous meta-analyses.

Objective: The objective was to quantify the effect of ≥3 g OBG/d on serum cholesterol concentrations in humans and investigate potential effect modifiers.

Design: A meta-analysis was performed on 28 RCTs comparing ≥3 g OBG/d with an appropriate control. Systematic searches were undertaken in PubMed, AGRICOLA, and Scopus between 1 January 1966 and 6 June 2013, plus in-house study reports at CreaNutrition AG. Studies were assessed with regard to inclusion/exclusion criteria, and data were extracted from included studies by reviewers working independently in pairs, reconciling differences by consensus. Estimates of the mean reduction in serum cholesterol from baseline between the OBG and control diets were analyzed by using random-effects meta-analysis models and meta-regression.

Results: OBG in doses of ≥3 g/d reduced low-density lipoprotein (LDL) and total cholesterol relative to control by 0.25 mmol/L (95% CI: 0.20, 0.30; P < 0.0001) and 0.30 mmol/L (95% CI: 0.24, 0.35; P < 0.0001), respectively, with some indication of heterogeneity (I^2 = 0.13 and P = 0.067). There was no significant effect of OBG on high-density lipoprotein (HDL) cholesterol or triglycerides and no evidence that dose (range across trials: 3.0–12.4 g/d) or duration of treatment (range: 2–12 wk) influenced the results. LDL cholesterol lowering was significantly greater with higher baseline LDL cholesterol. There was a significantly greater effect for both LDL and total cholesterol in subjects with diabetes compared with those without (although based on few studies).

Conclusions: Adding ≥3 g OBG/d to the diet reduces LDL and total cholesterol by 0.25 mmol/L and 0.30 mmol/L, respectively, without changing HDL cholesterol or triglycerides. Am J Clin Nutr 2014;100:1413–21.

Keywords β-glucan, HDL cholesterol, LDL cholesterol, total cholesterol, triglyceride, meta-analysis, oats

INTRODUCTION

European guidelines for the management of dyslipidemias (1) note the strong evidence supporting a relation between total and LDL cholesterol lowering and reduced cardiovascular disease risk and recommend that lifestyle modifications, such as consuming 5–15 g/d soluble fiber from oat products, may be beneficial. U.S. guidelines (2) recommend a multifaceted lifestyle approach, including dietary options for LDL cholesterol lowering, such as consuming 10–25 g/d soluble fiber from oat products and other foods.

Oat β-glucan (OBG), the main soluble fiber found in oats, is seen as the main active component responsible for their cholesterol-lowering effect. Health claims regarding the association between cholesterol lowering and soluble fiber from oat products/OBG have been approved by food standards agencies worldwide [United States: U.S. Food and Drug Administration (3); Canada: Health Canada (4); Europe: European Food Safety Authority (5); Australia and New Zealand: Food Standards Australia New Zealand (6); Malaysia: Ministry of Health Malaysia (7)]. Except for Malaysia, these approvals are based on a diet containing at least 3 g OBG/d, although specific conditions may vary; for example, the Food and Drug Administration (3) allows individual servings, including 0.75 g OBG, whereas the European Food Safety Authority (8) requires 1-g portions.

A meta-analysis of 12 randomized controlled trials (RCTs) published between 1985 and 1991 showed that oat products reduced total cholesterol by 0.13 mmol/L, with a higher intake of soluble fiber (≥3 compared with <3 g/d) having a significantly...
greater effect (9). A meta-analysis of 25 RCTs published between 1966 and 1996 showed that oat products reduced total and LDL cholesterol, respectively, by 0.040 and 0.037 mmol/L per gram of daily soluble fiber intake (10). Kelly et al. (11) included 8 RCTs published between 1991 and 2005 in a meta-analysis showing that oat products reduced total and LDL cholesterol, respectively, by 0.19 and 0.18 mmol/L.

However, an updated meta-analysis is timely and important for several reasons. The previous meta-analyses included studies with intakes of oat soluble fiber <3 g/d; thus, their results do not provide an accurate estimate of the effect on serum cholesterol of complying with the food standards agencies’ requirement that cholesterol-lowering claims for oats relate to daily intakes of ≥3 g oat soluble fiber. Also, we found 10 studies including ~1200 subjects published since 2005, which more than doubles the amount of data now available for meta-analysis. Finally, previous meta-analyses did not consider the molecular weight (MW) of OBG, but this may be important because high MW may be necessary to obtain a significant cholesterol-lowering effect. Therefore, our primary objective was to quantify the effect of consuming ≥3 g high-MW OBG/d on serum LDL, HDL, and total cholesterol and triglyceride concentrations by using data from RCTs comparing OBG with a control treatment.

METHODS

Literature search and study selection

Three electronic databases (PubMed, www.ncbi.nlm.nih.gov/pubmed; AGRICOLA, agricola.nal.usda.gov; and Scopus, www.elsevier.com/online-tools/scopus) were searched for relevant published articles (papers and abstracts) between 1 January 1966 and 6 June 2013. The reference lists of articles found to be relevant were checked for additional articles of relevance. The in-house collection of study reports at CreaNutrition AG was searched as well (CreaNutrition AG is the Switzerland-based international marketing, sales, and research subsidiary of Swedish Oat Fiber AB). The keywords used for the search were as follows: 1) (oat fibre OR oat fiber OR oat bran OR oat β-glucan OR oats OR oat bran concentrate OR oatmeal) AND cholesterol and 2) (oat fibre OR oat fiber OR oat bran OR oat β-glucan OR oats OR oat bran concentrate OR oatmeal) AND serum lipids. Searches were limited to human studies and publications in English.

An initial screening was undertaken for potentially relevant studies by 2 reviewers (EJB and ST). Articles on studies testing OBG as a dietary intervention for lipid lowering were retrieved and allocated a unique paper identification number. All retrieved studies were assessed with regard to inclusion and exclusion criteria and study quality (Supplemental Tables 1 and 2). Trials investigating the effect of ≥3 g OBG/d (MW ≥100 kDa) for a minimum of 2 wk were to be included.

The increased intake of OBG could be achieved by consumption of a range of food products such as bread, muesli, breakfast cereals, cereal bars, biscuits, cereal drinks, muffins, and powders that contained OBG in 1, 2, or more eating occasions during the day. The control treatment could consist of a comparable food product (bread, muesli, breakfast cereals, cereal bars, biscuits, cereal drinks, muffins, or powders) without soluble fiber (i.e., low in fiber) or a comparable food product high in insoluble fiber. We selected 2 wk as the minimum intervention period in keeping with a previous meta-analysis (10), because the cholesterol-lowering effect is not immediate.

Because the MW of OBG influences its cholesterol-lowering effect (12), we wanted to exclude studies with low MW OBG, which would not be expected to have much effect at a dose of 3 g/d. A MW of 100 kDa was chosen because this represents more than 95% degradation of native OBG (MW ~2000 kDa), and below 100 kDa OBG displays Newtonian behavior (no entanglements) and has a low viscosity at the relevant concentrations. Even though 3 g OBG/d with a MW of 210 kDa has been reported to have no significant effect on LDL cholesterol, because there is no standard procedure to extract and measure OBG MW, we wanted to err on the side of including studies that may not have used high enough MW rather than excluding those that did. However, studies did not have to report MW to be included. Studies not reporting MW could be included if one or more interventions were considered to contain high MW OBG, based on our knowledge of the effects of different food processing on β-glucan MW. Depolymerization is usually observed during production of OBG concentrates and extracts (13, 14). In general, the production of porridge, muesli, biscuits, cereal bars, and muffins does not greatly affect MW (15–17). Typical extrusion conditions are not severe enough to cause much degradation of β-glucan (12, 16, 17), but high shear can cause significant depolymerization (12).

The study populations included were generally healthy free-living normocholesterolemic or hypercholesterolemic adult men and women from the general population; subjects could be lean, overweight, or obese and could have type 2 diabetes. The quality criteria were based on Appendix H of the European Food Safety Authority guidance for the preparation and presentation of the application for authorization of a health claim (18). Assessors worked in pairs independently to complete assessments for the publications (EJB + ST, EJB + TMSW, ST + TMSW). They then discussed their results and reconciled differences by consensus.

Data extraction and statistical analysis

Assessors worked in pairs to extract data from all included studies into a data collection Excel spreadsheet (AW + EJB, ST, and TMSW). They then discussed their results and reconciled differences by consensus. The following information was extracted from each study whenever available: trial design (parallel group, crossover); number of subjects randomized per treatment arm; health status of study population (healthy, hyperlipidemic, diabetic); mean age; percentage of subjects who are male; details of OBG treatment and control treatment; type of control treatment (diet alone, cereal/low soluble fiber); daily dose of OBG (g/d total); background diet (standard diet or advice on weight maintenance, low fat/cholesterol lowering, energy restriction); treatment duration (wk); mean baseline LDL and total cholesterol (mmol/L); mean LDL, HDL, and total cholesterol and triglyceride concentrations with standard deviations of individual observations or standard errors of means, both at baseline and after treatment and, if available, for change from baseline for each treatment; and mean difference in the change from baseline between the OBG and the control diets with standard deviations of individual observations or standard errors of means.

For trials comparing one dose of OBG with a control treatment, an estimate of the mean difference in cholesterol between OBG
RESULTS

The initial screening yielded 355 publications. Based on titles and abstracts, 97 were reviewed for inclusion in the meta-analysis (Figure 1). Of these, 69 were excluded because they did not meet the inclusion and exclusion criteria. Twelve studies were excluded because the daily consumption of OBG was <3 g/d.

These low-dose interventions had the potential to misrepresent the magnitude of the effect of products carrying health claims and the associated health benefit. Of the 97 studies reviewed, information on MW was reported or found in related publications for 14 studies reporting 20 comparisons (some articles included several doses of OBG). Of these 20, 6 comparisons were excluded on this basis. Available data for included studies are given in Table 1. Depolymerization is typically observed during production of bread because of its susceptibility to enzymes during fermentation (41). One study (44) used a combination of bread and muffins, and another (32) used bread alone, but no MW data were provided, so they were included.

The characteristics of the 27 published (12, 16, 17, 23–46) and 1 unpublished (Y. Donazzolo, M. Latreille-Barbier, C. Ruel, S. Layre, R. Alken, M. Macmahon, unpublished results, 2006) RCTs included in the meta-analysis are described in Table 1. Eighteen RCTs had a parallel group design and 10 a crossover design. The dose of OBG ranged from 3.0 to 12.4 g/d, and...
## TABLE 1
Study characteristics\(^1\)

<table>
<thead>
<tr>
<th>Study (^2)</th>
<th>No. of subjects randomized</th>
<th>Duration of treatment, wk</th>
<th>Dose of OBG, g/d</th>
<th>Health status</th>
<th>Treated group</th>
<th>MW of OBG, kDa</th>
<th>Type of control treatment</th>
<th>Background diet</th>
<th>Male, %</th>
<th>Mean age, y</th>
<th>Mean baseline LDL-C, mmol/L</th>
<th>Mean baseline TC, mmol/L</th>
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<tbody>
<tr>
<td>Anderson (25)</td>
<td>21</td>
<td>3</td>
<td>12.4 H-chol</td>
<td>Out bran</td>
<td>1600-3000</td>
<td>Wheat bran</td>
<td>Standard</td>
<td>100</td>
<td>61.0</td>
<td>3.47</td>
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<tr>
<td>Beck 5-6 g (16)</td>
<td>46</td>
<td>12</td>
<td>5.5 Healthy</td>
<td>Out bran</td>
<td>1600-3000</td>
<td>Refined wheat/rice</td>
<td>Energy restr</td>
<td>0</td>
<td>37.4</td>
<td>3.04</td>
<td>5.10</td>
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<td>12</td>
<td>8.5 Healthy</td>
<td>Out bran</td>
<td>1600-3000</td>
<td>Refined wheat/rice</td>
<td>Energy restr</td>
<td>0</td>
<td>37.3</td>
<td>2.93</td>
<td>5.05</td>
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<td>Berg (26)</td>
<td>235</td>
<td>4</td>
<td>5.0 H-chol</td>
<td>Diet + oat bran</td>
<td>2100</td>
<td>Refined wheat/rice/corn</td>
<td>Weight adv</td>
<td>48</td>
<td>51.0</td>
<td>3.27</td>
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<td>3.2 H-chol</td>
<td>Out bran</td>
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<td>Standard</td>
<td>40</td>
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<td>Chen (27)</td>
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<td>12</td>
<td>7.3 Healthy</td>
<td>Out bran</td>
<td></td>
<td></td>
<td>Standard</td>
<td>20</td>
<td>52.4</td>
<td>3.79</td>
<td>5.48</td>
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<td>6</td>
<td>3.6 Healthy</td>
<td>Out bran</td>
<td></td>
<td></td>
<td>Farina</td>
<td>Low fat/chol</td>
<td>54</td>
<td>49.7</td>
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<td>6</td>
<td>4.0 Healthy</td>
<td>Out bran</td>
<td></td>
<td></td>
<td>Farina</td>
<td>Low fat/chol</td>
<td>71</td>
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<td>45</td>
<td>6</td>
<td>6.0 Healthy</td>
<td>Out bran</td>
<td></td>
<td></td>
<td>Farina</td>
<td>Low fat/chol</td>
<td>62</td>
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<td>Oatmeal + oat bran</td>
<td>900</td>
<td>Wheat Bran</td>
<td>Standard</td>
<td>100</td>
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<td>3.0 H-chol</td>
<td>Out bran</td>
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<td></td>
<td>900 Wheat Bran</td>
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<td>3.1 H-chol</td>
<td>Out bran</td>
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<td></td>
<td>Rice starch</td>
<td>Standard</td>
<td>52</td>
<td>51.7</td>
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<td>3.0 H-chol</td>
<td>Out bran</td>
<td></td>
<td></td>
<td>Corn flour</td>
<td>Low fat/chol</td>
<td>32</td>
<td>49.0</td>
<td>3.55</td>
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<td>Liatis (32)</td>
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<td>3</td>
<td>3.0 Diabetic</td>
<td>Out bran</td>
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<td></td>
<td>Refined wheat flour</td>
<td>Standard</td>
<td>56</td>
<td>63.0</td>
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<td>Miká (24)</td>
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<td>Out bran</td>
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<td></td>
<td>Refined wheat/corn</td>
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<td>Pins (33)</td>
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<td>5.4 Healthy</td>
<td>Oat bran + oat bran</td>
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<td>Refined wheat</td>
<td>Standard</td>
<td>51</td>
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<td>Queenan (34)</td>
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<td>6.0 H-chol</td>
<td>Out bran</td>
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<td></td>
<td>Dextrose</td>
<td>Standard</td>
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<td>44.9</td>
<td>4.15</td>
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<td>6</td>
<td>6.7 Healthy</td>
<td>Oatmeal</td>
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<td></td>
<td>Mixed refined grains</td>
<td>Energy restr</td>
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<td>44.6</td>
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<td>41</td>
<td>8</td>
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<td></td>
<td>Wheat bran</td>
<td>Low fat/chol</td>
<td>56</td>
<td>47.8</td>
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<td>Wolever 4L (12)</td>
<td>150</td>
<td>4</td>
<td>4.0 Healthy</td>
<td>Out bran</td>
<td>210</td>
<td>Wheat bran</td>
<td>Standard</td>
<td>39</td>
<td>52.4</td>
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<td>Out bran</td>
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<td>Wheat bran</td>
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<td>Out bran</td>
<td>850</td>
<td>Wheat bran</td>
<td>Standard</td>
<td>45</td>
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<td>Wolever 3H (12)</td>
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<td>Out bran</td>
<td>2210</td>
<td>Wheat bran</td>
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<td>46</td>
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<td>Zhang (37)</td>
<td>182</td>
<td>6</td>
<td>3.3 H-chol</td>
<td>Oatmeal</td>
<td></td>
<td></td>
<td>Wheat noodles</td>
<td>Weight adv</td>
<td>39</td>
<td>53.2</td>
<td>4.24</td>
<td>6.18</td>
</tr>
</tbody>
</table>

\(^1\) H-chol, hypercholesterolemic; LDL-C, LDL cholesterol; low fat/chol, low fat/cholesterol lowering; MW, molecular weight; OBG, oat β-glucan; restr, restricted; TC, total cholesterol; up, unpublished; Weight adv, weight maintenance advice.

\(^2\) For studies with more than one OBG arm, the number of subjects randomized equals those randomly allocated to the particular OBG arm plus those randomly allocated to the control arm. For Beck et al. (16), the 2 OBG doses were 5-6 and 8-9 g/d. For Davidson et al. (28), the 3 OBG doses were 84 g oatmeal (3.6 g/d), 56 g oat bran (4 g/d), and 84 g oat bran (6 g/d). For Wolever et al. (12), the 4 OBG doses were 4 g/d low MW (4L), 3 g/d medium MW (3M), 4 g/d medium MW (4M), and 3 g/d high-MW (3H).
treatment duration ranged from 2 to 12 wk. Twelve studies recruited healthy subjects, 13 recruited subjects with hypercholesterolemia, and 3 recruited subjects with type 2 diabetes. The study quality scores are provided in Supplemental Table 3.

The effect of OBG relative to control on LDL cholesterol for all studies and dose amounts is shown in Figure 2A. There was some indication of heterogeneity among the studies, although this did not reach statistical significance ($P = 0.13$). The random-effects analysis showed a statistically significant difference for OBG relative to control of $-0.25$ mmol/L (95% CI: $-0.30$, $-0.20$; $P < 0.0001$). There was evidence of an increasing effect with baseline LDL cholesterol ($P = 0.039$) and a significant, increased effect for subjects with diabetes relative to healthy subjects ($P = 0.013$, Supplemental Table 4 and Supplemental Figure 1).

There was an indication of an increasing effect with baseline total cholesterol ($P = 0.074$), age ($P = 0.080$), percentage of subjects who were male ($P = 0.054$), and for hypercholesterolemic subjects relative to healthy subjects ($P = 0.079$), although effects did not reach statistical significance. No other factors showed a significant effect.

The effect of OBG relative to control on total cholesterol for all studies and dose amounts is shown in Figure 2B. There was some indication of heterogeneity among the studies ($P = 0.067$). The random-effects meta-analysis showed a statistically significant difference for OBG relative to control of $-0.30$ mmol/L (95% CI: $-0.35$, $-0.24$; $P < 0.0001$). There was a significantly greater effect for subjects with type 2 diabetes relative to those without diabetes ($P = 0.004$, Supplemental Table 4). No other factor showed a significant effect.

Because of the influence of the Beck et al. (16) study on the HDL cholesterol results, the LDL and total cholesterol meta-analyses were repeated without this study. Omitting the Beck et al. study had little effect on the results for LDL and total cholesterol.

OBG tended to reduce serum triglyceride concentrations slightly compared with control, but the difference was not significant ($-0.023$ mmol/L; 95% CI: $-0.060$, $0.015$; $P = 0.23$, Figure 3B). There was little evidence of heterogeneity ($P = 0.58$).

To assess the effect of study quality on the results, we undertook meta-regressions on each of the dichotomized quality questions (Q11–Q14), as well as a composite of Q9a and Q9b, for LDL and total cholesterol. These indicated a reduction in the OBG effect between studies that met the quality criterion.

**FIGURE 2** Forest plots of estimates of mean differences (95% CIs). LDL cholesterol: heterogeneity ($x^2$, $P = 0.13$; $I^2 = 22\%$) (A). TC: heterogeneity ($x^2$, $P = 0.067$; $I^2 = 28\%$) (B). The area of the circle is proportional to the inverse variance of the estimate of mean difference. For studies with more than one OBG arm, the number of subjects randomized equals those randomly allocated to the particular OBG arm plus those to the control arm. For Beck et al. (16), the 2 OBG doses were 5–6 and 8–9 g/dL. For Davidson et al. (28), the 3 OBG doses were 84 g oatmeal (3.6 g/d), 56 g oat bran (4 g/d), and 84 g oat bran (6 g/d). For Wolever et al. (12), the 4 OBG doses were 4 g/d low MW (4L), 3 g/d medium MW (3M), 4 g/d medium MW (4M), and 3 g/d high MW (3H). diff, difference; MW, molecular weight; OBG, oat β-glucan; TC, total cholesterol; up, unpublished.
compared with those that did not (Supplemental Table 4). However, these differences were not statistically significant apart from Q13. The effect of OBG was significantly less by 0.11 mmol/L ($P = 0.024$) on LDL cholesterol and 0.13 mmol/L ($P = 0.021$) on total cholesterol for trials in which blinding of outcome assessors was specifically mentioned relative to the other trials. Larger effects of OBG on total cholesterol were associated with studies of a lower precision (i.e., larger 95% CI), although this was less obvious for LDL cholesterol.

**DISCUSSION**

We found that diets containing $\geq 3$ g OBG/d reduce serum total and LDL cholesterol relative to control by 0.30 and 0.25 mmol/L, respectively, with no effect on HDL cholesterol or triglycerides. Although generally confirming the results of previous meta-analyses that oat products reduce serum cholesterol, the present results differ in that the magnitude of the effects seen are 50–100% greater than those reported in previous meta-analyses. This is important because our study provides a more accurate assessment of the effect on serum cholesterol of following the recommendations of food standards agencies to consume $\geq 3$ g OBG/d compared to previous meta-analyses that included studies in which subjects consumed $<3$ g OBG/d and studies in which OBG was more than 95% degraded.

We found no significant effect of dose or duration of treatment of the range of OBG doses (3.0–12.4 g/d) and durations of treatment (2–12 wk) used in the RCTs included in the meta-analysis. The former suggests that a cholesterol-lowering effect of oats can be achieved with the minimum dose (3 g/d) considered effective by regulatory agencies and that consuming more may not have any additional effect. The lack of dose response is curious and somewhat unexpected because a significant effect of dose was found in both previous meta-analyses that tested for it (9, 10). However, the previous meta-analyses included studies with intakes of OBG, 3 g/d, and one (10) also assumed a linear dose-response relation. Also, the ability of our meta-analysis to detect an effect of dose may be limited because of unknown effects of unmeasured confounding variables, such as the source of oats or the nature of the food products used to deliver OBG, which may have affected the physicochemical properties and, hence, viscosity of OBG (47). It is believed that the cholesterol-lowering effect of OBG depends on its viscosity in the small intestine, which, in turn, is affected by the MW and amount of $\beta$-glucan in solution. The MW of $\beta$-glucan may be reduced by $\beta$-glucanase before being incorporated into food products by exposure to $\beta$-glucanase naturally present in foods to which it is added (e.g., wheat flour) (41) or by heat and pressure exerted on foods during processing. Viscosity is inversely related to log (MW). Wolever et al. (12) showed a reduced cholesterol-lowering effect of OBG with low MW (210 kDa) relative to a medium MW (530 kDa) or high MW (2210 kDa). We restricted our meta-analysis to studies including OBG with MW $\geq 100$ kDa. However, because there is no standardized method for measuring the MW of $\beta$-glucan, this could have been

![FIGURE 3](https://academic.oup.com/ajcn/article-abstract/100/6/1413/4576477/1418-WHITEHEAD-ET-AL)

**FIGURE 3** Forest plots of estimates of mean difference (95% CIs). HDL cholesterol: heterogeneity ($\chi^2$, $P < 0.001$; $I^2 = 81\%$) (A). Triglycerides: heterogeneity ($\chi^2$, $P = 0.58$; $I^2 = 0\%$) (B). The area of the circle is proportional to the inverse variance of the estimate of mean difference. For studies with more than one OBG arm, the number of subjects randomized equals those randomly allocated to the particular OBG arm plus those to the control arm. For Beck et al. (16), the 2 OBG doses were 5–6 and 8–9 g/d. For Davidson et al. (28), the 3 OBG doses were 84 g oatmeal (3.6 g/d), 56 g oat bran (4 g/d), and 84 g oat bran (6 g/d). For Wolever et al. (12), the 4 OBG doses were 4 g/d low MW (4L), 3 g/d medium MW (3M), 4 g/d medium MW (4M), and 3 g/d high MW (3H). diff, difference; MW, molecular weight; OBG, oat $\beta$-glucan; up, unpublished.
a source of confounding in our analysis. The amount of β-glucan in solution in the small intestine depends on its ability to be solubilized and released from the food matrix (i.e., bioaccessibility); β-glucan solubility is known to be reduced by low water availability in a food and storage of hydrated β-glucan at cool temperatures. Because we had no way of assessing the bioaccessibility of the β-glucan in most studies included in our analysis, it also is a potential source of confounding.

The lack of effect of study duration on the results suggests that the effect of oats on serum cholesterol is durable, as found by Bazzano et al. (48), but none of the studies we included lasted for longer than 12 wk.

We found evidence that the LDL cholesterol-lowering effect of oats was greater in subjects with type 2 diabetes and subjects with higher baseline LDL cholesterol. Previous studies suggest that the cholesterol-lowering effect of OBG is greater in nonwhites than in whites (24, 31, 49); this, taken together with our results suggesting that OBG has a greater effect in subjects with type 2 diabetes, might indicate that the OBG reduces serum cholesterol via a mechanism or mechanisms related in some way to dysglycemia, insulin resistance, and/or insulin secretion. Although type 2 diabetes generally is not associated with increased LDL cholesterol, it is associated with increased secretion of VLDL particles, which, after interaction with lipoprotein-lipase, hepatic-lipase, and cholesterol-ester transfer protein, eventually are metabolized to become LDL particles. In addition, this suggests that those with increased risk for cardiovascular disease due to high cholesterol or diabetes will obtain at least as much, if not more, benefit from the cholesterol-lowering effect of oats as individuals without these risk factors; however, these conclusions are based on a limited number of studies and should be interpreted with caution.

The studies included a wide range of subjects, including healthy individuals and those with hypercholesterolemia and type 2 diabetes. The studies were conducted in Europe, North America, Asia, and Australia. A wide range of common food products were used to study the effect of β-glucan, including rolled oats, whole oat flour, oat bran, bread, muffins, muesli, breakfast cereals, cereal bars, and biscuits. Therefore, it would appear that the results are applicable to the general population and that benefits could be achieved eating regularly consumed foods.

The funnel plots indicated that the studies with a lower precision had a tendency to show a more beneficial effect of OBG on total cholesterol, although this effect was less obvious for LDL cholesterol. This is sometimes taken to infer that there is publication bias—that is, a number of small studies have been undertaken and not published because the effect is not statistically significant or is negative. However, there may be other reasons, such as larger studies being undertaken after promising results from small studies. As suggested by the meta-regressions on the quality questions, the funnel plots show some association between OBG effect and study quality, with lower study quality being associated with a larger effect. However, this might be explained by study quality improving over time.

We conclude that there is robust evidence that consuming oats or oat-containing food products containing at least 3 g OBG/d with MW ≥100 kDa reduces serum cholesterol in lean, overweight, or obese male and female adults without diabetes and those with type 2 diabetes. Directions for future research include the need for high-quality studies to determine whether there is a dose-response effect of OBG and whether the effect persists in the long term (i.e., longer than 3 mo). Because the purported mechanisms of cholesterol lowering require a significant viscosity of the gastrointestinal contents, and this is supported by recent work (12, 41), future studies need to consider the physicochemical properties of β-glucan in food products targeted at lowering total and LDL cholesterol to ensure the intended physiologic effects.
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