# **REVIEW ARTICLE**



# The long-term effects of primary school-based obesity prevention interventions in children: A systematic review and meta-analysis

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# **Summary**

**Introduction:** This systematic review and meta-analysis investigate the long-term effects of primary school-based obesity prevention interventions on body-mass index (and z-scores), waist circumference (and z-scores) and weight status.

Methods: Four databases were searched for studies from date of inception until June 8th, 2021. We included randomized controlled trials (RCT) and non-RCTs investigating effects ≥12 months post-intervention of primary school-based interventions with intervention duration ≥6 months and containing a diet and/or physical activity component on outcomes of interest. Articles were assessed on risk of bias and methodological quality by RoB2 and ROBINS-I. Meta-analysis was performed and results were narratively summarized. Evidence quality was assessed with GRADE.

**Results:** Nineteen studies were included, 9 were pooled in a meta-analysis. No long-term effects were found on body-mass index ( $+0.06 \text{ kg/m}^2$ ; CI95% = -0.38, 0.50;  $I^2 = 66\%$ ), body-mass index z-scores (-0.08; CI95% = -0.20, 0.04;  $I^2 = 36\%$ ), and waist circumference (+0.57 cm; CI95% = -0.62, 1.75;  $I^2 = 13\%$ ). Non-pooled studies reported mixed findings regarding long-term effects on body-mass index, body-mass index z-scores and weight status, and no effects on waist circumference and waist circumference z-scores. Evidence certainty was moderate to very low.

**Discussion:** No clear evidence regarding long-term effects of primary school-based interventions on obesity-related outcomes was found. Recommendations for further research and policy are discussed.

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# KEYWORDS

elementary school, prevention, program, lifestyle, sustained effect

Abbreviations: BMI, body-mass index; CI, confidence interval; GRADE, grading of recommendations assessment, development and evaluation; PRISMA-P, preferred reporting items for systematic reviews and meta-analysis protocol; PROSPERO, prospective register of systematic reviews; RCT, randomized controlled trial; RoB2, risk of bias 2; ROBINS-I, risk of bias in non-randomized studies of interventions; WC, waist circumference; WHO, World Health Organization; zBMI, BMI-for-age z-scores; zWC, WC-for-age z-scores.

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# 1 | INTRODUCTION

Current estimates by the World Health Organization (WHO) are that almost one in five children worldwide are affected by overweight or obesity. Although a recent stabilization has been confirmed regarding the overweight and obesity prevalence in higher-income countries, the prevalence overall remains high.<sup>2,3</sup> Studies have shown that effective interventions for obesity prevention accomplish more positive effects on body-mass index (BMI) for younger children.<sup>4</sup> A recent Cochrane review by Brown and colleagues (2019) counted 153 randomized controlled trials (RCT) investigating the effectiveness of interventions that include diet or physical activity components, or both, designed for preventing overweight and obesity in children and adolescents.<sup>5</sup> They have concluded that interventions that focus only on physical activity can reduce the risk of obesity for children aged 6 to 12 years old, whereas they found no evidence for the effectiveness of diet interventions and some evidence for the effectiveness of interventions with a combined diet component and physical activity component.<sup>5</sup> The authors could not entirely explain the differences in the effects of interventions by the setting or duration of the interventions.<sup>5</sup> Although obesity prevention interventions may be implemented in many settings, schools have been suggested to be the optimal place to deliver interventions due to continuous contact with children<sup>6</sup> and reaching children with a wide range of different backgrounds. Additionally, the results of other systematic reviews support the promising role of school-based interventions, 8,9 specifically primary school-based interventions. 10 for the prevention of childhood obesity.

RCTs are often considered the golden standard for effectiveness evaluation studies. 11 However, only a limited number of RCTs have a long-term follow-up (≥12 months post-intervention). It might be challenging to conduct such long-term RCTs due to funding issues or because it is unethical to withhold children from effective interventions for more years. In their review, Brown and colleagues (2019) encourage researchers to perform more long-term follow-up studies into the sustained effects of interventions. If RCTs are not feasible, researchers might turn to other types of designs, like natural experiments, quasi-experimental and observational studies with comparison groups as these can also generate causal evidence with good external validity. 12-14 Insights in the long-term effects of interventions are especially relevant for the transition from childhood into adolescence as the WHO highlighted that adolescence is a period of excessive weight gain because adolescents gain more freedom in food choices, while their physical activity levels often decrease.15

To the best of our knowledge, a recent systematic review focusing on the long-term effects of primary school-based obesity prevention interventions in children is not available. Such a review aggregating the available evidence may either help policymakers in developing and implementing successful interventions of which the effects sustain into adolescence and adulthood or may highlight the need for supplementary interventions at later ages. Therefore the research question in this systematic review and meta-analysis is:

What are the long-term effects (≥12 months follow-up post-intervention) of primary school-based obesity prevention interventions in children on multiple indicators of overweight and obesity? Intervention effects on BMI, BMI-for-age z-scores (zBMI), WC, WC-for-age z-scores (zWC) and weight status will be included as important measures of obesity and in predicting obesity-related health risk, as clinical guidelines suggest. <sup>16</sup>

## 2 | METHODS

#### 2.1 | Protocol

A protocol for this systematic review and meta-analysis was developed in line with the Preferred Reporting Items for Systematic reviews and Meta-analysis Protocol (PRISMA-P).<sup>17</sup> This protocol was prospectively registered within Prospective Register of Systematic Reviews (PROSPERO) on 02-04-2021 with registration ID: CRD 42021240446.

#### 2.2 | Inclusion and exclusion criteria

#### 2.2.1 | Type of primary studies

We included RCTs and studies with other controlled experimental and observational designs in this review. We excluded studies with prepost designs without control group.

# 2.2.2 | Intervention and setting

This review focusses on primary school-based obesity prevention interventions. Therefore, interventions had to be at least partially primary school-based to be included, but we allowed interventions in multiple settings. Following Brown and colleagues (2019), we included interventions containing at least a diet or physical activity component. Interventions could also include both components or other components in addition to diet or physical activity.

Only interventions with a duration of ≥6 months were included in this review because, according to the theory of Prochaska and DiClemente, a duration of at least 6 months is needed for behaviour change to stabilize when it involves physical activity. To study long-term effects, the intervention effects should have been studied following a period in which there was no exposure to the intervention. Thus, we only included primary studies in which intervention effects were studied ≥12 months post-intervention.

# 2.2.3 | Control

We included studies with a control or comparator condition without an intervention.



# 2.2.4 | Participants in intervention

As prospectively determined, we included studies in which at least 90% of the children were 6- to 12-year-olds at the start of the intervention. When we were not able to determine this criterion, all children in the study had to be within the age range for the study to be included. Furthermore, studies mentioning primary school-aged children, without the notion of an age range, were also considered eligible for inclusion.

# 2.2.5 | Participants at follow-up

Since this review focuses on long-term effects of interventions (≥12 months follow-up post-intervention), children studied at follow-up may have transcended the age of 12 years old. No criterion was set on age range at follow-up.

#### 2.2.6 | Context

In this review primary studies regarding the general population were included. Therefore, children with overweight or obesity were allowed as a part of the participant sample. However, studies that were solely focusing on children with overweight or obesity were excluded since this review focusses on interventions preventing these conditions in the general population. Studies with participants with other clinical conditions were also not eligible for inclusion in this review.

We applied no further restrictions regarding subgroups of the population (e.g., country of origin or socioeconomic status). Furthermore, no restrictions were drawn upon the country of origin of the studies.

#### 2.2.7 | Outcomes

Outcomes of interest for this review were BMI, zBMI, WC, zWC, and weight status. zBMI is a score in which BMI is adjusted to age and gender of the individual based on growth standards and zWC is a score in which WC is adjusted to age and gender.<sup>19</sup> Weight status can be categorized as having underweight, normal weight, overweight or obesity using common cut-off standards.

# 2.2.8 | Language

Studies in English were eligible for inclusion in this review. Studies in other languages were excluded due to language barriers of the authors.

# 2.3 | Search strategy

A systematic literature search without language restrictions or date limits was conducted to find relevant studies in four databases (Embase.com, Medline ALL Ovid, Web of Science SCI-EXPANDED & SSCI and Cochrane CENTRAL register of Trials) from date of inception until the 8th of June 2021. In addition, we hand-searched references of included studies and the Cochrane review by Brown and colleagues.<sup>5</sup>

Corresponding authors were contacted when the full texts were unavailable on the internet. Combined thesaurus terms and terms in title/abstract were used for three elements: intervention or program, body-mass index or weight status or overweight or obesity or waist circumference and children. The syntax and thesaurus terms within this search strategy were adapted to fit to each of the used databases. The full search is presented in supplemental file 1 (Appendix S1).

# 2.4 | Selection process

EndNote X9 Software was used to guide and organize the selection process of the studies that were found. Duplicates of records were retrieved and removed by guidance of Bramer and colleagues.<sup>20</sup> Hereafter title and abstracts were screened for eligibility by two independent reviewers (M.S. and M.B.). Then these studies were screened on full text to select the studies eligible for inclusion (M.S. and M.B.). A prospectively discussed order of reasons for exclusion was determined by the screening authors. Studies that were excluded based on full text screening, were categorized based on the reason of exclusion. When differences or disagreements between the two reviewers were encountered in the process, discussion with a third reviewer (W.J.) took place to reach consensus.

#### 2.5 | Data extraction

All data was extracted to a standardized data extraction Excel sheet by two independent authors (M.S. and M.B.). Discrepancies were discussed until consensus was reached. If available, the following data was extracted for all included studies: (1) first author, (2) year of publication, (3) full reference, (4) study objective, (5) theoretical framework behind the intervention, (6) country of intervention, (7) funding sources, (8) conflict of interest statement, (9) study design, (10) intervention and control group characteristics (sample size, age, gender), (11) number of clusters, (12) numbers of participating primary schools, (13) intervention characteristics (duration, components and setting), (14) comparison or control condition, (15) follow-up period, (16) context, (17) outcomes at baseline, (18) outcomes at follow-up, (19) estimate of the effect sizes for the association between primary school-based intervention and outcome (coefficients, Cl's, *p*-values) and (20) retention rates.

Specifically for the meta-analysis, mean and standard deviations were extracted for the outcome variables. When results from multiple models were available, results from the model with the most confounders added were extracted. When multiple follow-up time points were available for a study we extracted data for all follow-up time

points. For the purpose of the meta-analyses, we chose the time point of the longest follow-up within adolescence for inclusion in the meta-analysis, as suggested by the Cochrane Handbook.<sup>21</sup> The remaining earlier follow-up time points were included in an additional sensitivity meta-analysis.

#### 2.6 Risk of bias assessment

A risk of bias assessment was performed by two independent authors (M.S. and M.B.) to assess the quality of each included study. When differences or disagreements between the two reviewers were encountered in the process, discussion with a third reviewer (W.J.) took place to reach consensus. The revised Cochrane Collaboration's Risk of Bias 2 (RoB2) tool was used for assessing risk of bias for randomized trials<sup>22</sup> and the Risk of Bias in Non-randomized studies of interventions (ROBINS-I) tool was used for assessing risk of bias for non-randomized studies.<sup>23</sup> The RoB2 assesses bias (1) in the randomization process, (2) due to deviations from intended interventions, (3) due to missing outcome data, (4) due to measurement of outcomes and, (5) in selection of the reported result. All types of bias were rated as low risk, some concerns or high risk. The overall risk of bias score was determined by the highest risk score. The ROBINS-I contains items for bias (1) due to confounding, (2) in selection of participants into the study, (3) in classification of interventions, (4) due to deviations from intended interventions, (5) due to missing data, (6) in measurement of outcomes and, (7) in selection of the reported result. All domains were rated as having low, moderate, serious or critical risk of bias. Like in the ROB2, the overall risk of bias score was determined by the highest risk score.

# 2.7 Data synthesis and analysis

The characteristics for all the included studies and interventions were described. We conducted a meta-analysis for intervention effects for which at least three studies were available with the same study design and outcome measure and if the outcome measures could be calculated into mean and standard deviation values per group at follow-up. We used the mean differences at follow-up in outcome measures between the intervention group and control group. We used randomeffects models to account for the between-study variance. The metaanalyses were conducted in Review Manager (version 5.4). The overall pooled mean difference and 95% confidence intervals (CI) were calculated. Forest plots were created to present the meta-analysis results. I<sup>2</sup> measures have been presented as an indication for the proportion of total variability due to between study heterogeneity. Additional sensitivity analyses were performed including available earlier followup time points. The remaining extracted results that could not be pooled in the meta-analyses or sensitivity meta-analyses were narratively described. Therefore, studies could be considered as both pooled for some pooled outcome variables and non-pooled for its other non-pooled outcome variables. The narratively described results were divided into results derived from RCT studies and results derived from non-RCT studies.

Summary of findings tables were created with the Grading of Recommendations Assessment, Development and Evaluation (GRADE) pro software to present the evidence quality across studies and certainty of effects of the interventions.<sup>24</sup> The evidence quality was assessed according to the GRADE handbook.<sup>25</sup> The effects that were included in the meta-analyses were added in the summary of findings tables as pooled outcome variables. Results that could not be included in the meta-analysis were narratively described in the summary of findings tables as the non-pooled outcomes. For the narratively described non-pooled outcomes a subdivision was created for evidence resulting from RCT studies and evidence resulting from non-RCT studies. The GRADE approach for rating the evidence quality began with rating the study design. The evidence quality rating could then be reduced by rating the factors: risk of bias, inconsistency in results, indirectness of evidence, imprecision, and publication bias. The evidence quality rating could be increased by the factors: large magnitude of effect, plausible confounding and dose-response gradient. Possible final grades of evidence were high, moderate, low, or very low certainty.

# 3 | RESULTS

#### 3.1 | Study selection

The process of inclusion and exclusion of articles during article selection is presented in a PRISMA flow chart in Figure 1.<sup>17</sup> A total of 15 887 records were identified through comprehensive database searches. After the removal of duplicates, 9188 records remained for screening. After screening on title and abstract, 211 articles remained for full text screening. From these 211 articles, a total of 192 articles were excluded. Most important reasons for exclusion were the study design (n=27), follow-up length (n=113) and age of the population (n=23). Supplemental file 2 (Appendix S1) presents a complete overview of the excluded references during full text screening per exclusion category. Eventually, 19 articles were found eligible for inclusion into this systematic review.<sup>26-44</sup> Of these 19 articles, 9 could be included into either the meta-analysis or additional sensitivity meta-analysis on earlier time-points.<sup>26,27,32,35-38,43,44</sup>

# 3.2 | Study characteristics

The most important study characteristics are presented in Table 1. Study objective, theory behind the intervention, scale-up, funding, conflict of interest statement, intervention content and control content are presented in Table S1. Furthermore, Table S3 presents a complete detailed overview of the results that were extracted from the 19 included articles in this review. Of the 19 studies, 16 were published in or after 2010. 26-31,33-36,38-41,43,44 The 19 studies were performed in 11 different countries, subdivided over the continents Europe



# **PRISMA 2009 Flow Diagram**

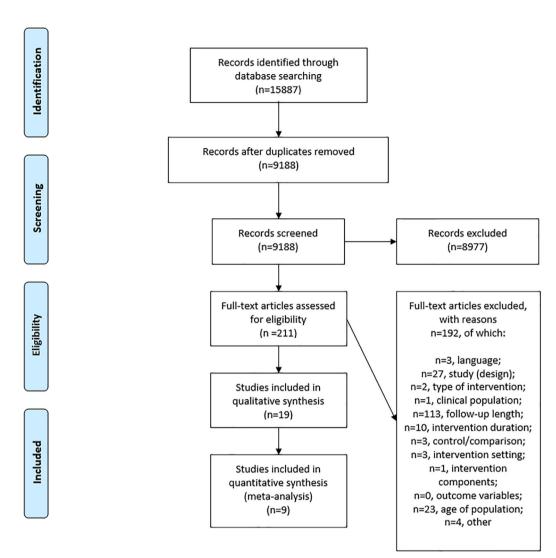


FIGURE 1 Flow diagram showing the selection process in this review

(10/19), <sup>26,27,30,32,33,35,36,41,43,44</sup> North-America (4/19), <sup>29,37,39,40</sup> Asia (3/19)<sup>28,31,34</sup> and Oceania (2/19). <sup>38,42</sup> Five studies reported to be based on the Social Cognitive Theory, <sup>26,30,31,40,42</sup> one reported the socioecological conceptual model, <sup>36</sup> one reported the Behaviour Change Theory, <sup>37</sup> one reported the concepts of the WHO Ottawa Charter for Health Promotion, <sup>39</sup> one reported Piaget's Cognitive Theory <sup>40</sup> and one reported the Behavioural Choice Theory. <sup>42</sup> Of the 19 studies, five interventions were considered a scale-up. <sup>26,29,31,34,37</sup> Most studies (13/19) were (cluster-)RCTs. <sup>26–28,31,32,34–38,42–44</sup> Other designs were a repeated measures crossover, <sup>29</sup> a non-randomized controlled trial, <sup>30</sup> two quasi-experimental, <sup>33,40</sup> a repeated cross-sectional <sup>39</sup> and a population-based prospective controlled <sup>41</sup> design. Four studies had a sample size between 100 and 250 participants, <sup>29,38,40,41</sup> six studies had a sample size between 251 and 500 participants, <sup>27,30,32,33,36,42</sup> five

studies had a sample size between 501 and 1000 participants  $^{28,31,35,39,44}$  and four studies had a sample size between 1001 and 5000 participants.  $^{26,34,37,43}$ 

All interventions took place in a primary school setting, <sup>26-44</sup> although six interventions additionally contained a home setting component <sup>28,35-37,42,44</sup> and one intervention additionally contained a home setting, community setting and online setting component. <sup>30</sup> Classroom or school teachers (partially) delivered the interventions in most studies. <sup>26,28-35,37,41</sup> Three interventions solely contained a diet component <sup>27,32,43</sup> and seven interventions solely contained an physical activity component, <sup>29,33,34,36,38,41,42</sup> whereas nine interventions contained both a diet and an physical activity component. <sup>26,28,30,31,35,37,39,40,44</sup> Eight interventions contained an additional non-diet non-physical activity component. <sup>26,28,35,37,39,40,42,44</sup> which is

TABLE 1 Characteristics of included studies

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| Author and year                  | Intervention/<br>study name   | Country                     | Study design  | Settings                                  | No of primary<br>schools No of<br>clusters in<br>analysis         | Intervention<br>delivered by                        | Intervention<br>components: Diet,<br>Physical activity,<br>Other      | Intervention<br>duration                     |
|----------------------------------|---|-----------------------------|---|---|---|---|---|--|
| Anderson, 2016 <sup>27</sup>     | The Active for Life<br>Year 5 (AFLY5)   | England                     | Cluster-randomized<br>controlled trial                | Primary school                            | 60 primary<br>schools<br>60 clusters                              | School teachers,<br>Learning support<br>assistants  | Diet, Physical activity,<br>Sedentary<br>behaviour, Screen<br>viewing | 6-7 months                                   |
|                                  | The Fruits and<br>Vegetables Make<br>the Marks (FVMM)<br>project  | Norway                      | Cluster-randomized controlled trial                   | Primary school                            | 38 primary<br>schools<br>38 clusters                              | Z<br>Z  | Diet  | 9 months                                     |
|                                  | The Family-Individual-<br>School (FIS)-based<br>comprehensive<br>childhood-obesity<br>intervention                | China                       | Cluster-randomized controlled trial                   | Primary<br>school,<br>Home                | 14 primary<br>schools<br>Clusters at class<br>level; amount<br>NR | Health teachers,<br>Teachers                        | Diet, Physical activity   | 34 months                                    |
|                                  | The Dance Dance<br>Revolution-based<br>exercise<br>intervention   | United States<br>of America | Repeated-measures<br>crossover design                 | Primary school                            | 1 primary school<br>Clusters NR                                   | Classroom teachers,<br>Research assistants          | Physical activity   | 7 months                                     |
|                                  | The GreatFun2Run<br>intervention  | England                     | Non-randomized<br>controlled trial                    | Primary<br>school,<br>Community,<br>Home, | 8 clusters  | School teachers                                     | Diet, Physical activity   | 10 months                                    |
| Habib-Mourad, 2020 <sup>32</sup> | The Ajyal Salima<br>("Healthy Kids")<br>intervention,<br>originating from the<br>"Health-E-PALS<br>intervention". | Lebanon                     | Stratified cluster-<br>randomized<br>controlled trial | Primary school                            | 36 primary<br>schools<br>36 clusters                              | Teachers, Researchers<br>(to train the<br>teachers) | Diet, Physical activity   | 16 months<br>(over two<br>academic<br>years) |
|                                  | The Christchurch obesity prevention programme in schools (CHOPPS)   | England                     | Cluster-randomized<br>controlled trial                | Primary school                            | 6 primary schools<br>29 clusters (class<br>level)                 | One investigator,<br>School teachers                | Diet  | 12 months                                    |
|                                  | The enhanced PE programme   | Slovenia                    | Quasi-experimental                                    | Primary school                            | 9 primary schools<br>Clusters NR                                  | General teacher,<br>Specialist PE teacher           | Physical activity   | 48 months                                    |
|                                  | The Happy 10 program  | China                       | Cluster-randomized<br>controlled trial                | Primary school                            | 20 primary<br>schools<br>132 clusters<br>(class level)            | Teachers, Health<br>educators                       | Physical activity   | 1 school year                                |
| Llaurado, 2018 <sup>36</sup>     |   | Spain                       |   |   |   |   |   | 28 months                                    |

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| Intervention<br>duration   |  | 12 months   | 36 months   | 9 months                            | 36 months                                 | 24 months   | 6-8 years   | 9 months   | 9 months   | 28 months                            |
|--|--|---|---|-------------------------------------|---|---|---|--|--|--------------------------------------|
| Intervention<br>components: Diet,<br>Physical activity,<br>Other | Diet, Physical activity,<br>Sedentary<br>behaviour                                 | Physical activity                                     | Diet, Physical activity,<br>Cigarette smoking   | Physical activity                   | Diet, Physical activity,<br>Mental health | Diet, Physical activity,<br>Screen viewing                              | Physical activity   | Physical activity, Sedentary behaviour, Screen viewing | Diet   |                                      |
| Intervention<br>delivered by                                     | School teachers,<br>university students<br>who acted as health<br>promoting agents | PE teachers   | Classroom teachers,<br>Food service<br>personnel, PE<br>specialists   | Instructors                         | School health<br>facilitators             | Local resource<br>teachers  | Regular school<br>teachers                                  | PE teachers,<br>Interventionists                       | Ϋ́<br>Z  |                                      |
| No of primary<br>schools No of<br>clusters in<br>analysis        | 38 primary<br>schools<br>Clusters NR   | 15 primary<br>schools<br>28 clusters (class<br>level) | 96 primary<br>schools<br>Clusters NR  | 2 primary schools<br>2 clusters     | 173 primary<br>schools<br>Clusters NR     | Primary schools<br>NR (1 school<br>district)<br>Clusters NR             | 4 primary schools<br>Clusters NR                            | 3 primary schools<br>17 clusters (class<br>level)      | 38 primary<br>schools<br>Clusters NR                             | 38 primary<br>schools                |
| Settings   | Primary<br>school,<br>Home   | Primary<br>school,<br>Home                            | Primary<br>school,<br>Home  | Primary school                      | Primary school                            | Primary school  | Primary school  | Primary<br>school,<br>Home                             | Primary school   |                                      |
| Study design   | Randomized<br>controlled trial   | Cluster-randomized<br>controlled trial                | Randomized<br>controlled trial  | Cluster-controlled randomized trial | Repeated cross-<br>sectional              | Quasi-experimental  | Population-based prospective controlled intervention study  | Cluster randomized controlled trial                    | Cluster-randomized<br>controlled trial                           | Cluster- randomized controlled trial |
| Country  |  | Switzer- land   | United States of America  | Australia                           | Canada                                    | United States<br>of America   | Sweden  | Australia  | Norway   | Spain                                |
| Intervention/<br>study name                                      | The Educació en<br>Alimentació (EDAl)<br>program                                   | The Kinder- und<br>Jugendsportstudie<br>(KISS)        | The Child and<br>Adolescent Trial for<br>Cardiovascular<br>Health (CATCH)<br>school and family<br>intervention<br>(CATCH III) | The CAPO kids trial                 | APPLE schools                             | The Integrated<br>Nutrition and<br>Physical Activity<br>Program (INPAP) | The Paediatric<br>Osteoporosis<br>Prevention (POP)<br>study | The Switch-Play intervention                           | The Fruits and<br>Vegetables Make<br>the Marks (FVMM)<br>Project |                                      |
| Author and year  |  | Meyer, 2014 <sup>37</sup>                             | Nader, 1999 <sup>38</sup>   | Nogueira, 2017 <sup>39</sup>        | Ofosu, 2018 <sup>40</sup>                 | Puma, 2013 <sup>41</sup>  | Rosengren, 2021 <sup>42</sup>                               | Salmon, 2008 <sup>43</sup>                             | Stea, 2018 <sup>44</sup>   | Tarro, 2014 <sup>45</sup>            |

(Continues)

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| Author and year                 | Intervention/<br>study name                      | Country Study design  | Settings   | No of primary<br>schools No of<br>clusters in Intervention<br>analysis delivered by                                     | Intervention<br>delivered by                                   | Intervention<br>components: Diet,<br>Physical activity,<br>Other   | Intervention<br>duration          |
|---------------------------------|--|---|--|---|--|--|-----------------------------------|
|                                 | The Educació en<br>Alimentació (EDAl)<br>program | EDAI)   | Primary 2<br>school,<br>Home   | 2 clusters Univer<br>who<br>pron  | University students<br>who acted as health<br>promoting agents | Diet, Physical activity,<br>Sedentary<br>behaviour   |                                   |
| Author<br>and year              | Follow-up length                                 | No. of total group participants at follow-up (% female) No. of intervention group participants at follow-up (% female) No. of control group participants at follow-up (% female)  | Baseline age total group (mean(SD))  Baseline age intervention group (mean(SD))  Baseline age control group (mean(SD)) | Follow-up age total group (mean(SD)) Follow-up age intervention group (mean(SD)) Follow-up age control group (mean(SD)) |  | Other specific<br>population<br>characteristics  | Retention<br>rate at<br>follow-up |
| Anderson,<br>2016 <sup>27</sup> | 12 months  | n=2221 (50.8%) in total $n=1064$ (48.9%) in intervention group $n=1157$ (52.5%) in control group  | 9.5 (0.3) y in total group<br>9.5 (0.3) y in<br>intervention group<br>9.5 (0.3) y in control group                     | NR in total group NR in intervention group NR in control group  |  | Z.   | 80.7%                             |
| Bere, 2014 <sup>28</sup>        | 36 and<br>84 months                              | n=320 (62%) in total $n=112$ (59%) in intervention group $n=208$ (64%) in control group   | 11.8 (NR) y in total group NR in intervention group NR in control group  | NR in total group  NR in intervention group  NR in control group  |  | At baseline, 45% of intervention group was in 7th grade. The remainder in 6th grade. At baseline, 48% of control group was in 7th grade. The remainder in 6th grade.   | 16.4%                             |
| Cao, 2019 <sup>29</sup>         | 24 and 36 months                                 | n = 807 (46.5%) in total $n = 406 (42.9%)$ in intervention group $n = 401 (50.1%)$ in control group   | NR in total group 7.00 (0.41) y in intervention group 6.82 (0.24) y in control group                                   | NR in total group 13.68 (0.40) y in intervention group 13.50 (0.24) y in control group                                  |  | Z.   | 33.0%                             |
| Gao, 2013 <sup>30</sup>         | 12 months  | <ul> <li>n = 156 (41.2%) in total, including 53 participants with no longterm effects measured due to continued intervention in year 2.</li> <li>n = 68 (NR) in intervention group</li> <li>n = 44 (NR) in control group</li> </ul> | 10.32 (0.91) y in total group  Between 10 and 12 y in intervention group  Between 10 and 12 y in control group         | 10.28 (0.90) y in total group  NR in intervention group  NR in control group  |  | The participating school had the highest obesity rate in the state of Utah, US. Vast majority of children were from economically disadvantaged, Latino immigrant families.  Approximately 73% of children received free lunch. | 79.3%                             |

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| Retention<br>rate at<br>follow-up  | 71.5%   | 65.1%  | 67.4%   | 72.8%   | 87.7%  | 21.4%   |
|--|---|--|---|---|--|---|
| Other specific<br>population<br>characteristics  | The majority of participants were of white British ethnicity (intervention 94.8% and control 96.5%) | The study included public schools (primarily low socioeconomic status) and private schools (primarily middle and high socioeconomic status). | Ϋ́Z   | Urban locations   | Z Z  | ж<br>Z  |
| Follow-up age total group (mean(SD)) Follow-up age intervention group (mean(SD)) Follow-up age control group (mean(SD))  | NR in total group<br>NR in intervention group<br>NR in control group                                | NR in total group NR in intervention group NR in control group   | 11.6 (NR) y, with a range of 10.0–13.9 y in total group NR in intervention group NR in control group  | NR in total group  NR in intervention group  NR in control group                              | NR in total group  NR in intervention group  NR in control group                                       | NR in total group<br>15.6 (0.53) y in intervention<br>group         |
| Baseline age total group (mean(SD)) Baseline age intervention group (mean(SD)) Baseline age control group (mean(SD))   | NR in total group<br>8.8 (NR) y in intervention group<br>8.9 (NR) y in control group                | 9.95 (1.13) y in total group<br>9.81 (0.68) y in intervention<br>group<br>10.13 (0.68) y in control group                                    | 8.6 (NR) y, with a range of 7.0–10.9 y in total group 8.7 (0.8) y for boys in intervention group 8.7 (1.0) y for girls in intervention group 8.6 (0.9) y for boys in control group 8.7 (0.9) y for girls in control group 8.7 (0.9) y for girls in control group 8.7 (0.9) y for girls in control group | NR in total group 7.76 (0.33) y in intervention group 7.71 (0.32) y in control group          | 9.3 (0.7) y in total group<br>NR in intervention group<br>NR in control group                          | NR in total group (age range of 7–8 y).<br>NR in intervention group |
| No. of total group participants at follow-up (% female) No. of intervention group participants at follow-up (% female) No. of control group participants at follow-up (% female) | n=421 (NR) in total $n=206$ (NR) in intervention group $n=215$ (NR) in control group                | n=806 (NR) in total $n=457$ (NR) in intervention group $n=349$ (NR) in control group   | $n=434 \ (48.2\%)$ in total $n=219 \ (45.2\%)$ in intervention group $n=215 \ (51.2\%)$ in control group $n=215 \ (51.2\%)$ in control group  | n=324 (48.8%) in total $n=160$ (46.3%) in intervention group $n=164$ (51.2%) in control group | n = 4120 (48.6%) in total $n = 2028 (46.4%)$ in intervention group $n = 2092 (50.7%)$ in control group | n=503 (NR) in total $n=154$ (NR) in intervention group              |
| Follow-up length   | Between 18 and 20 months  | 12 months  | 24 months   | 84 months   | 12 months  | 48 months   |
| Author<br>and year   | Gorely, 2011 <sup>31</sup>  | Habib-Mourad,<br>2020 <sup>32</sup>  | James, 2007 <sup>33</sup>   | Jurak, 2013 <sup>34</sup>   | Li, 2010 <sup>35</sup>   | Llaurado,<br>2018 <sup>36</sup>                                     |

(Continues)

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|  | No. of total group participants at follow-up (% female) No. of intervention group                       | Baseline age total group<br>(mean(SD))   | Follow-up age total group<br>(mean(SD))   |   |                                   |
|--|---|--|---|---|-----------------------------------|
| participants at follow-up (% female) No. of control group particit at follow-up (% female)           | participants at follow-up (%<br>female)<br>No. of control group participants<br>at follow-up (% female) | Baseline age intervention group (mean(5D)) Baseline age control group (mean(5D))                         | Follow-up age intervention group (mean(SD)) Follow-up age control group (mean(SD))                        | Other specific<br>population<br>characteristics                               | Retention<br>rate at<br>follow-up |
| n=349 (NR) in control group  | control group   | NR in control group  | 14.9 (0.36) y in control group  |   |                                   |
| n=173 (50.3%) first graders in total $n=116$ (61.2%) fifth graders in total                          | rst graders in<br>1.2%) fifth   | NR in total group  | NR in total group   | First graders: 31% overweight in intervention group and                       | 57.6%                             |
| n = 108 (47%) first graders in intervention group $n = 81 (59%)$ fifth graders in intervention group | raders in<br>aders in   | 6.9 (0.3) y for first graders in intervention group 10.9 (0.5) y for fifth graders in intervention group | 10.6 (0.3) y for first graders in intervention group 15.0 (0.5) y for fifth graders in intervention group | 14% overweight in control group at baseline. Fifth graders: 20% overweight in |                                   |
| n = 65 (55%) first graders in control group $n = 35$ (66%) fifth graders in control group            | ars in<br>ers in  | 6.9 (0.3) y for first graders in control group 11.1 (0.6) y for fifth graders in control group           | 10.6 (0.4) y for first graders in control group<br>15.1 (0.6) y for fifth graders in control group        | both groups at<br>baseline. 14–31%<br>migrants.                               |                                   |
| n=3660 (NR) in total   |   | 8.74 (NR) y in total group   | NR in total group   | Ethnically diverse  | 72.7%                             |
| n=2164 (NR) in intervention group  | ention  | NR in intervention group   | NR in intervention group  |   |                                   |
| n=1496 (NR) in control group   | group   | NR in control group  | NR in control group   |   |                                   |
| n = 240 (43.8%) in total   |   | 10.6 (0.6) y in total group  | 12.3 (0.6) y in total group   | NR  | 70.8%                             |
| n=155~(42.6%) in interventi group  | ention  | 10.5 (0.6) y in intervention group   | 12.2 (1.0) y in intervention group  |   |                                   |
| n=85  (44.8%) in control group   | ol group  | 10.7 (0.6) y in control group  | 12.3 (0.6) y in control group   |   |                                   |
| n=540 (NR) in total  |   | NR in total group  | NR in total group   | From low socioeconomic  | ΑN                                |
| n = 202 (48.3%) in intervention group  | vention   | 10.8 (0.4) y in intervention group   | 13.8 (1.4) y in intervention group  | neighbourhoods  |                                   |
| n=338~(59.1%) in control group   | rol group   | 10.9 (0.4) y in control group  | 14.0 (1.3) y in control group   |   |                                   |
| n=191 (NR) in total  |   | NR in total group  | NR in total group   | 2nd graders at baseline;  | ΑN                                |
| n=91(NR) in intervention group   | ention group  | NR in intervention group   | NR in intervention group  | 8th graders at follow-  |                                   |
| n=100 (NR) in control group  | rol group   | NR in control group  | NR in control group   | children (i.e., 62%) in<br>free school lunch                                  |                                   |

| Continued) |  |
|------------|--|
| TABLE 1    |  |

|   | No. of total group participants at follow-up (% female) No. of intervention group                                 | Baseline age total group<br>(mean(SD))   | Follow-up age total group<br>(mean(SD))  |  |                                   |
|---|---|--|--|--|-----------------------------------|
| Follow-up length                          | participants at follow-up (% female)  No. of control group participants at follow-up (% female)                   | Baseline age intervention group<br>(mean(SD))<br>Baseline age control group<br>(mean(SD))                | Follow-up age intervention<br>group (mean(SD))<br>Follow-up age control group<br>(mean(SD))                | Other specific<br>population<br>characteristics        | Retention<br>rate at<br>follow-up |
|   |   |  |  | program and 59.1% was Hispanic.                        |                                   |
| Between 36 and 60 months (mean 48 months) | n=124~(46.8%) in total $n=81~(44.4%)$ in intervention group   | NR in total group 7.5 (0.6) y for boys in intervention group 7.5 (0.6) y for girls in intervention group | NR in total group 18.8 (0.2) y for boys in intervention group 18.7 (0.4) y for girls in intervention group | Elementary schools with a uniform socioeconomic status | 35.5%                             |
|   | n=43(51.2%) in control group  | 8.0 (0.7) y for boys in control group<br>7.9 (0.6) y for girls in control group                          | 18.8 (0.3) y for boys in control group<br>18.7 (0.3) y for girls in control group                          |  |                                   |
| 12 months                                 | n=268 (NR) in total   | 10.67 (0.42) $y^a$ for boys in total group 10.67 (0.33) $y^a$ for girls in total group                   | NR in total group  | Low socioeconomic status                               | 90.8%                             |
|   | n=84 (NR) in BM/FMS intervention group $n=60$ (NR) in BM intervention group $n=69$ (NR) in FMS intervention group | NR in intervention group   | NR in intervention group   |  |                                   |
|   | n=55 (NR) in control group  | NR in control group  | NR in control group  |  |                                   |
| 168 months                                | n=1081 (53.5%) in total $n=322$ (53.0%) in intervention group   | NR in total group<br>NR in intervention group  | 26.5 (NR) y in total group<br>26.5 (0.53) y in intervention<br>group                                       | 10 to 12 year olds at baseline.                        | 52.8%                             |
|   | n=759~(55.6%) in control group  | NR in control group  | 26.5 (0.51) y in control group   |  |                                   |
| 24 months                                 | n=619 (NR) in total   | NR in total group  | NR in total group  | 7 to 8 year olds at                                    | 26.3%                             |
|   | $\it n=198$ (NR) in intervention group  | NR in intervention group   | 12.51 (0.52) y in intervention group   | baseline.  |                                   |
|   | n=421 (NR) in control group   | NR in control group  | 11.88 (0.35) y in control group  |  |                                   |

Abbreviations: BM, behavioural modification; FMS, fundamental movement skills; NA, not applicable; NR, not reported; PE, physical education; SD, standard deviation; US, United States; y, years. 
<sup>a</sup>Reported in months, but calculated into years by us.

TABLE 2 Quality assessment for RCTs with the RoB2 tool

|                                  | Randomization process | Item 2 Deviations from intended interventions | Item 3<br>Missing<br>outcome<br>data | Item 4 Measurement of the outcome | Item 5<br>Selection of the<br>reported<br>result | Overall score |
|----------------------------------|-----------------------|---|--------------------------------------|-----------------------------------|--|---------------|
| Anderson, 2016 <sup>27</sup>     | Low risk              | High risk                                     | Low risk                             | Low risk                          | High risk  | High risk     |
| Bere, 2014 <sup>28</sup>         | High risk             | Some concerns                                 | Some concerns                        | High risk                         | Some concerns                                    | High risk     |
| Cao, 2019 <sup>29</sup>          | Some concerns         | Some concerns                                 | Some concerns                        | Low risk                          | Some concerns                                    | Some concerns |
| Habib-Mourad, 2020 <sup>32</sup> | Some concerns         | High risk                                     | Low risk                             | Low risk                          | Some concerns                                    | High risk     |
| James, 2007 <sup>33</sup>        | Low risk              | Some concerns                                 | High risk                            | Low risk                          | Some concerns                                    | High risk     |
| Li, 2010 <sup>35</sup>           | Low risk              | High risk                                     | Low risk                             | Low risk                          | Some concerns                                    | High risk     |
| Llaurado, 2018 <sup>36</sup>     | Low risk              | Some concerns                                 | Low risk                             | Low risk                          | Low risk   | Some concerns |
| Meyer, 2014 <sup>37</sup>        | Low risk              | Some concerns                                 | Low risk                             | Low risk                          | Low risk   | Some concerns |
| Nader, 1999 <sup>38</sup>        | Low risk              | Some concerns                                 | Low risk                             | Low risk                          | Low risk   | Some concerns |
| Nogueira, 2017 <sup>39</sup>     | Low risk              | Some concerns                                 | High risk                            | Low risk                          | Some concerns                                    | High risk     |
| Salmon, 2008 <sup>43</sup>       | Low risk              | Some concerns                                 | Some concerns                        | Low risk                          | Some concerns                                    | Some concerns |
| Stea, 2018 <sup>44</sup>         | Some concerns         | High risk                                     | High risk                            | High risk                         | Some concerns                                    | High risk     |
| Tarro, 2014 <sup>45</sup>        | Some concerns         | Some concerns                                 | Low risk                             | Some<br>concerns                  | Low risk   | Some concerns |

further specified in Table 1. Ten interventions had a duration between 6 and 12 months, <sup>26,27,29,30,32,34,36,38,42,43</sup> seven had a duration between 13 and 36 months <sup>28,31,35,37,39,40,44</sup> and two had a duration of >36 months. <sup>33,41</sup> Seven studies investigated long-term effects at a follow-up of 12 months post-intervention, <sup>26,28,29,31,34,38,42</sup> seven studies investigated effects at a follow-up between 13 and 36 months post-intervention, <sup>27,28,30,32,36,37,44</sup> four studies investigated effects at a follow-up between 37 and 60 months post-intervention <sup>35,39-41</sup> and three studies investigated effects at a follow-up of >60 months post-intervention. <sup>27,33,43</sup>

# 3.3 | Risk of bias

The risk of bias assessment for the 13 RCTs with the RoB2 tool is reported in Table 2. The overall risk of bias was assessed as "high risk" in seven RCTs and as "some concerns" in six RCTs. The largest risk of bias in RCTs was due to deviations from intended interventions, being classified as high risk in 4/13 RCTs and as some concerns in 9/13 RCTs. The two main concerns with this bias were that deviations from the intended intervention could have affected outcomes and the absence of an intention to treat analysis. Risk of bias due to the randomization process mainly came forth from absence of information on both the randomization method and method used to hide the allocation sequence from participants. Risk of bias due to missing outcome data mainly came forth from missing outcome data not being equally distributed over groups. Finally risk of bias due to selection of the reported result came forth from studies lacking any pre-specified study protocol registration.

The risk of bias assessment for the six non-RCTs with the ROBINS-I tool is reported in Table 3. Among the six non-RCTs the

overall risk of bias was assessed as "critical risk" in one non-RCT and as "serious risk" in five non-RCTs. The risk of bias in non-RCTs was largest with regard to confounding (mainly due to not controlling for baseline differences), classification of interventions (mainly due to lack of defining control group) and missing data (mainly due to the unequal distribution of missing data over groups). For all six non-RCTs no information was available to rate the domain of selection of the reported result, forthcoming from studies lacking a prospectively registered study protocol.

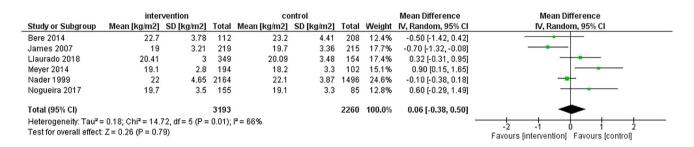
# 3.4 | Meta-analyses on long-term intervention effects

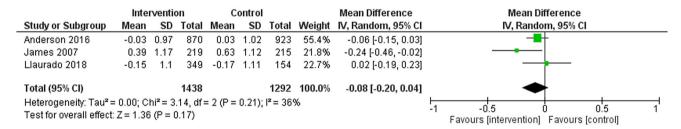
Comparisons were possible between intervention and control groups for BMI, zBMI and WC (Figure 2). Table 4 contains a summary of findings which presents the evidence quality and certainty for these pooled comparisons. The three funnel plots for these meta-analyses are in Figure S1, Figure S2 and Figure S3 respectively. In these meta-analyses the time points of the longest follow-up<sup>27,35</sup> within adolescence were included when there were multiple follow-up time points<sup>27,35,44</sup> of the same study available. A complete overview of the GRADE rating assessments per GRADE item is presented in Table S2.

Results from six studies (n=5453 participants) that were included in the meta-analysis for the evaluation of the long-term effect of the interventions on BMI show that the interventions had no effect on BMI in primary school-aged children  $\ge 12$  months post-intervention.  $^{27,32,35-38}$  The mean difference in BMI between intervention and control groups was 0.06 kg/m² (CI95% = -0.38, 0.50; n=5453, Z=0.260, p=0.79).  $I^2$  was indicated as 66% ( $\tau^2=0.18$ ,  $\chi^2=14.72$ ,

 TABLE 3
 Quality assessment for non-RCTs with the ROBINS-I tool

|                                  | Item 1 Confounding | Item 2<br>Selection of<br>participants<br>into the<br>study | Item 3  Classification of interventions | Item 4 Deviations from intended interventions | Item 5 Missing data | Measurement of outcomes | Item 7<br>Selection<br>of the<br>reported<br>result | Overall score |
|----------------------------------|--------------------|---|---|---|---------------------|-------------------------|---|---------------|
| Gao, 2013 <sup>30</sup>          | Serious risk       | Low risk  | Low risk                                | Low risk                                      | NI                  | Low risk                | No information                                      | Serious risk  |
| Gorely, 2011 <sup>31</sup>       | Moderate<br>risk   | Low risk  | Low risk                                | Low risk                                      | Serious<br>risk     | Serious risk            | No information                                      | Serious risk  |
| Jurak, 2013 <sup>34</sup>        | Serious risk       | Low risk  | Low risk                                | Low risk                                      | Serious<br>risk     | Low risk                | No information                                      | Serious risk  |
| Ofosu, 2018 <sup>40</sup>        | Moderate<br>risk   | Serious risk  | Serious risk                            | Low risk                                      | NI                  | Low risk                | No information                                      | Serious risk  |
| Puma, 2013 <sup>41</sup>         | Critical risk      | Low risk  | Serious risk                            | Low risk                                      | NI                  | Low risk                | No information                                      | Critical risk |
| Rosengren,<br>2021 <sup>42</sup> | Serious risk       | Low risk  | Low risk                                | Low risk                                      | Moderate<br>risk    | Low risk                | No information                                      | Serious risk  |





|   | Inter     | vention |        | Co             | ntrol   |       |        | Mean Difference     |           | Mean Difference                          |   |
|---|-----------|---------|--------|----------------|---------|-------|--------|---------------------|-----------|--|---|
| Study or Subgroup   | Mean [cm] | SD [cm] | Total  | Mean [cm]      | SD [cm] | Total | Weight | IV, Random, 95% CI  |           | IV, Random, 95% CI                       |   |
| Llaurado 2018   | 72.06     | 8.44    | 349    | 71.33          | 9.72    | 154   | 38.3%  | 0.73 [-1.04, 2.50]  |           |  |   |
| Meyer 2014  | 64.7      | 7.3     | 189    | 63.3           | 7.9     | 102   | 35.5%  | 1.40 [-0.45, 3.25]  |           | +  |   |
| Nogueira 2017   | 65.3      | 8       | 155    | 66.1           | 8.5     | 85    | 26.2%  | -0.80 [-3.00, 1.40] |           |  |   |
| Total (95% CI)  |           |         | 693    |                |         | 341   | 100.0% | 0.57 [-0.62, 1.75]  |           |  |   |
| Heterogeneity: Tau <sup>2</sup> =<br>Test for overall effect: |           |         | (P = 0 | .32); I² = 13% |         |       |        |                     | <u>-4</u> | -2 0 2                                   | 4 |
|   |           | /       |        |                |         |       |        |                     |           | Favours [intervention] Favours [control] |   |

**FIGURE 2** Three random-effects meta-analysis of the mean difference in baseline to follow-up change in respectively BMI, zBMI and waist circumference between intervention and control groups

df = 5, p = 0.01). The mean difference estimate for BMI has a very low evidence certainty according to the GRADE assessment due to risk of bias and imprecision. Results from three studies (n = 2730 participants) that were included in the meta-analysis on zBMI show that the interventions had no long-term effect on zBMI at  $\geq$ 12 months of follow-up in primary school-aged children. The mean difference in zBMI between intervention and control groups was -0.08 (CI95% = -0.20, 0.04; n = 2730, Z = 1.360, p = 0.17).  $I^2$  was indicated as 36% ( $\tau^2$  = 0.00,  $\chi^2$  = 3.14, df = 2, p = 0.21). Certainty of the

evidence for the long-term effect estimate of the interventions on zBMI is very low. Results from three studies (n=1034 participants) that were included in the meta-analysis on WC show that the interventions had no long-term effect on WC at follow-up in primary school-aged children. The mean difference in WC between intervention and control groups was 0.57 cm (CI95% = -0.62, 1.75; n=1034, Z=0.940, p=0.35).  $I^2$  was indicated as 13% ( $\tau^2=0.14$ ,  $\chi^2=2.29$ , df = 2, p=0.32). Certainty of the evidence for the long-term effect estimate of the interventions on WC is low.

# 3.5 | Sensitivity meta-analyses on long-term intervention effects

In the sensitivity analyses studies with shorter follow-up time-points<sup>27,44</sup> were used instead of longest follow-up time-points.<sup>27,35</sup> The forest plot and funnel plot of the additional sensitivity meta-analysis for BMI, in which the 84 months follow-up results<sup>27</sup> have been replaced by the 36 months follow-up results,<sup>27</sup> are presented in Figure S4 and Figure S7 respectively.

The forest plots and funnel plots of the additional sensitivity meta-analyses for zBMI and WC, in which the study by Llaurado and colleagues (2018)<sup>35</sup> has been replaced by the study by Tarro and colleagues (2014),<sup>44</sup> are presented in Figure S5, Figure S6, Figure S8 and Figure S9.

The results from the three additional sensitivity meta-analyses correspond to those of the three meta-analyses. The mean difference in BMI between intervention and control groups was 0.09 kg/m<sup>2</sup> (CI95% = -0.33, 0.50; n = 5453, Z = 0.410, p = 0.68). <sup>27,32,35-38</sup> The

TABLE 4 Summary of Findings for the sustained effect of primary school-based obesity prevention interventions in children aged 6-12 years

Patient or population: Primary school-aged children aged 6 to 12 years old

Setting: At least one component of the intervention was primary school-based

Intervention: Obesity prevention interventions with a diet component, physical activity component or both components.

|  |                                      |                             |                                     |                                    |                           |  | Anticipated absolute effects   |  |  |
|--|--------------------------------------|-----------------------------|-------------------------------------|------------------------------------|---------------------------|--|--|--|--|
| Pooled outcomes                                    | No. of participa<br>(studies) Follow |                             | Certainty of t                      |                                    | Relative effe<br>(95% CI) | ct   | Risk with control group  | Risk difference with intervention group*   |  |
| ВМІ  | 5453 (6 RCTs)                        |                             | ⊕○○○<br>Very low <sup>a</sup>       |                                    | -                         |  | The mean BMI at follow-up was <b>20,55 kg/m</b> <sup>2</sup>   | MD <b>0.06 higher</b> (CI -0.38, 0.50)   |  |
| zBMI   | 2730 (3 RCTs)                        |                             | ⊕○○○<br>Very low <sup>a</sup>       |                                    | -                         |  | The mean zBMI at follow-up was <b>0.11</b>   | MD <b>0.08 lower</b> (CI -0.20 0.04)   |  |
| WC   | 1034 (3 RCTs)                        |                             | $\bigoplus_{Low^b}\bigcirc\bigcirc$ |                                    | -                         |  | The mean WC at follow-up was <b>67,62 cm</b>   | MD <b>0.57 higher</b><br>(CI -0.62, 1.75)  |  |
| Not-pooled   | outcomes                             | No. of part<br>(studies) Fo | •                                   | Certainty of evidence (            |                           | Effe   | cts narrative <sup>#</sup>   |  |  |
| ВМІ  |                                      | 5820 (3 RC                  | ^^^^^                               | ⊕⊕○○<br>^^^^^^<br>⊕○○○<br>Very low | .^^^^                     | for for in AAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAA | ed findings were reported by thround no association between ground that increase in BMI since betweention group compared with horself and the findings were reported by through the found no association between the findings that the difference tervention group and control group to the first the fi | up and BMI. The third study asseline was lower in the the control group.  ANNANANANANANANANA  ee non-RCT studies. Two reen group and BMI. The gramme* year interaction are in BMI between the pup decreased more and |  |
| BMI adjusted for the age-<br>sex-population median |                                      | 268 (1 RCT)                 |                                     | Low <sup>b</sup> s<br>ii<br>t      |                           | se<br>in<br>th                                 | study reported that the change<br>ex-population median was not sign<br>tervention group and FMS inter-<br>be control group. However chango<br>pulation median was significant<br>M/FMS intervention group com  | gnificantly lower for the BM<br>vention group compared to<br>ge in BMI minus sex-age<br>ly lower for the combined  |  |
| zBMI   |                                      | ^^^^^                       |                                     | Very low <sup>a</sup>              |                           | reinno   | dixed findings were reported by three RCT studies. One study reported significant lower zBMI at 24 and 36 months follow-up in the intervention group compared with the control group for normal weighted children at baseline. A second study reported a significantly larger decrease in zBMI in the intervention group compared with the control group. A third study found no association between group and zBMI.   |  |  |
| WC 4   |                                      |                             |                                     | ⊕⊕⊖⊖<br>Low <sup>d</sup>           | ) One                     |  | e RCT study found no association between group and WC.   |  |  |
| zWC  |                                      | 2655 (2 RC                  | CTs)                                | ⊕⊖⊖⊖<br>Very low                   | <i>r</i> <sup>a</sup>     |  | associations between group and und in two RCT studies.   | zWC or change in zWC we  |  |

TABLE 4 (Continued)

| Not-pooled outcomes | No. of participants (studies) Follow-up | Certainty of the evidence (GRADE) <sup>1</sup> | Effects narrative#  |
|---------------------|---|--|---|
| Weight status       | 7059 (9 RCTs) ^^^^^^^ 896 (3 non-RCT)   | Overy low <sup>a</sup>                         | Nine RCT studies reported mixed findings. Six studies did not report significant associations between group and weight status. A seventh study reported a significantly lower percentage of overweight participants in the intervention group compared with the control group at 84 months follow-up, whereas no significant association was found at 36 months follow-up. An eighth study found a significant association between intervention group and lower overweight percentage at 36 months follow-up. The ninth study found a significant association between the combined BM/FMS intervention group and decrease in overweight percentage compared with the control group, but no significant association was found between the BM intervention group or FMS intervention group and change in overweight percentage compared with the control group. |

Note: # = narrative effects for the not-pooled outcomes are based on the results in Table S4. **GRADE Working Group grades of evidence** High certainty: we are very confident that the true effect lies close to that of the estimate of the effect. Moderate certainty: we are moderately confident in the effect estimate: the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different. Low certainty: our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect. Very low certainty: we have very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of effect.

Grading explanations: 1. Inconsistency component of GRADE is uncertain for the not-pooled outcomes. No available sufficient information on heterogeneity or confidence intervals. 2. Full explanations and reasons underlying downgrading are presented supplemental file 5 (Appendix S1). a. 2 levels downgraded for risk of bias and 1 level downgraded for imprecision. b. 1 level downgraded for risk of bias and 1 level downgraded for imprecision. c. 1 level downgraded for risk of bias. d. 2 levels downgraded for risk of bias.

Abbreviations: BM, behavioural modification; BMI, body-mass index; CI, confidence interval; FMS, fundamental movement skills; MD, mean difference; RCT, randomized controlled trial; WC, waist circumference.

\* The risk in the intervention group (and its 95% confidence interval) is based on the assumed risk in the comparison group and the relative effect of the intervention (and its 95% CI).

mean difference in zBMI between intervention and control groups was  $-0.08 \text{ kg/m}^2$  (CI95% = -0.19, 0.02; n=2846, Z=1.550, p=0.12). The mean difference in WC between intervention and control groups was 0.62 cm (CI95% = -0.51, 1.75; n=1150, Z=1.080, p=0.28). 36,38,44

# 3.6 | Long-term effectiveness of interventions on non-pooled outcomes

The long-term intervention effects on the non-pooled outcomes are narratively described in Table 4a subdivision is made between evidence originating from RCTs and evidence originating from non-RCTs. A complete overview of the GRADE rating assessments per GRADE item, and accompanied with our explanations for grading up and down, is presented in Table S2.

Six studies, of which three RCTs and three non-RCTs, studied intervention effects on BMI and findings were mixed. Of the three RCTs, two studies found no association between intervention and a long-term impact on BMI with follow-up periods of 24 and 168 months. The third RCT found a lower BMI increase at 12 months of follow-up since baseline in the intervention group

compared with the control group.<sup>34</sup> The certainty of the evidence across the RCTs following the GRADE assessment was rated as low. Of the three non-RCTs, two studies found no association between intervention and a long impact on BMI with follow-up periods of 20 and 48 months.<sup>30,41</sup> One study found a significant interaction effect between intervention and time. In this study the differences in BMI between groups decreased toward the null during the follow-up period, which suggests that the intervention effect did sustain up to 84 months of follow-up.<sup>33</sup> The certainty of the evidence across the non-RCTs following the GRADE assessment was rated as very low.

Only one non-RCT reported on the change since baseline in BMI adjusted for the age-sex-population median. This study had three intervention arms and no significant effect on the outcome was found for the intervention group with solely a behavioural modification component and for the intervention group with solely a fundamental movement skills component in comparison with the control group. The intervention group with combined behavioural modification and functional movements skills components however was associated with a significant lower BMI adjusted for the age-sex-population median in comparison with the control group. Certainty of the evidence was rated as low.

Regarding zBMI, three RCTs reported mixed findings. One study found no association between intervention and zBMI at 12 months follow-up.<sup>31</sup> One study found significant lower zBMI values in the intervention group compared with the control group at 24 and 36 months of follow-up.<sup>28</sup> Another study reported a significantly larger decrease in zBMI from baseline to 12 months of follow-up in the intervention group compared with the control group.<sup>34</sup> For zBMI the certainty of the evidence across RCTs following the GRADE assessment was rated as moderate. Furthermore, one non-RCT found no association between intervention and zBMI at 20 months of follow-up.<sup>30</sup> Certainty of the evidence following the GRADE assessment was rated as very low.

One study examined the intervention effect on WC and found no association between obesity prevention interventions and WC at 20 months of follow-up. <sup>30</sup> The certainty of this evidence was rated as low. In addition, two studies reported on intervention effects on zWC at respectively 12 and 24 months of follow-up and both studies found no association between group and zWC or change in zWC. <sup>26,32</sup> The certainty of the evidence was rated as very low.

A total of 12 studies reported on the effect of obesity prevention interventions on weight status, of which nine RCTs and three non-RCTs. Of the nine RCTs, the majority of six studies found no association between the intervention and weight status. 26,31,32,35,43,44 In one RCT a lower percentage of overweight was found in the intervention group in comparison with the control group at 84 months of followup, whereas this association between group and percentage of overweight was not apparent at 36 months of follow-up.<sup>27</sup> Another RCT found a similar association at 24 and 36 months of follow-up.<sup>28</sup> Last, one RCT with 12 months follow-up found a significant association between the combined behavioural modification and functional movement skills intervention group and decrease in the percentage of children with overweight compared with the control group, but no association was found between the behavioural modification intervention group or functional movement intervention group and change in overweight percentage compared with the control group.<sup>42</sup> The certainty of the evidence across RCTs was rated as very low. Furthermore, three non-RCTs found no association between obesity prevention interventions and weight status ranging between 12 and 60 months of follow-up. 29,39,40

# 4 | DISCUSSION

We included 19 studies in this review, of which 9 RCT studies were included in either the pooled meta-analyses or pooled sensitivity meta-analyses. Sixteen studies were used for non-pooled analyses. The results from the meta-analyses suggest no long-term effects of obesity prevention interventions on the pooled outcome WC, whereas the evidence is very uncertain about the long-term effects on BMI and zBMI. The additional sensitivity meta-analyses, including the earlier follow-up time points, revealed similar results. Narrative results have been subdivided in evidence originating from RCTs and evidence resulting from non-RCTs. Beneficial long-term effects on the

non-pooled outcomes were found in only a limited number of studies, irrespective of study design. Summarizing, the results of this review do not show clear evidence regarding the long-term effects of primary school-based obesity prevention interventions in children on obesity-related outcomes

In total five studies reported positive long-term intervention effects on non-pooled outcomes<sup>27,28,33,34,42</sup> and one study in the meta-analyses reported positive long-term intervention effects.<sup>32</sup> When having a closer look at these intervention studies (Table S3), they show much variation in characteristics such as intervention components, intervention duration and follow-up length. Likewise, the other 13 studies that found no evidence for a long-term effect ≥12 months post-intervention in any outcome were very diverse in terms of those characteristics. Furthermore, there is also much variability regarding additional settings on top of the primary schoolbased setting and regarding the theoretical framework underlying the intervention in both the studies for which long-term effects were found and the studies for which no long-term effects were found. As we could not statistically perform subgroup analysis due to lack of power, the role of intervention duration, intervention components, follow-up length, setting and theoretical framework on long-term intervention effectiveness remains uncertain. More targeted evidence in this area is needed and may be gathered using innovative study designs in future studies and by using predictive modelling combining intervention characteristics and individual participant data.<sup>45</sup>

The interpretation of studied long-term intervention effects above needs consideration of the immediate post-intervention effects as well. Some interventions might have been effective to start with and others not. We have been able to retrieve this information for 15 of the 19 intervention studies in both the included studies in this review<sup>26,28-39,42,44</sup> and in earlier published studies.<sup>46-52</sup> For 4 of these 15 studies, positive long-term effects were found. 28,32,34,42 Our findings indicate that the intervention effects sustained up to 12 months post-intervention in two studies<sup>34,42</sup> and have sustained and have been improving up to 36 and 24 months since immediately post-intervention in the other two studies.<sup>28,32,46</sup> When looking into the 11 studies for which no long-term effects on any outcome were found in this review, 26,29-31,33,35-39,44 three interventions were not effective immediately post-intervention as well on any outcome, 29,31,37,47 whereas eight interventions showed effectiveness on at least one outcome immediately post-intervention.  $^{26,30,33,35,36,38,39,44,48-52}$  The lack of sustained and long-term intervention effects in most of the studies may have several causes, but the obesogenic environment that surrounds and shapes the health behaviour of children and adolescents certainly plays an important role, 53,54 opposing the effects that an ended intervention might have accomplished. However, methodological issues of the included studies might give alternative explanations for the lack of sustained and longterm intervention effects. Sample sizes in general were small, as 15 of the 19 studies had a sample size between 100 and 1000 participants. Attrition of participants at follow-up was apparent in all the included studies, and ranged between 16.4% and 87.7%. More specifically, the retention rates were even lower for the studies with a longer followup length. Such participant attrition possibly results in power issues to adequately assess intervention effects at long-term follow-up.

This review has been aggregating the available evidence on the long-term effects of primary school-based obesity preventive interventions and intends to help the research field on long-term intervention effect studies forward. The findings in this review must be interpreted with caution due to a few limitations. Based on the GRADE assessment tool, the certainty of the evidence for the pooled and non-pooled outcomes was generally low to very low, mainly due to the relatively high risks of bias across studies. This is in line with our quality assessment score for the included studies. As mentioned above sample sizes and attrition in included studies might have influenced the lack of positive findings. Furthermore, the small sample sizes in combination with the low number of included studies influenced the possibilities for interpretation of heterogeneity in the metaanalyses. 55,56 As we included a small number of studies in each metaanalysis, the confidence intervals were wide. This causes uncertainty regarding the systemacy of the  $l^2$ -measure. A visual exploration of the forest plots suggests some additional undetected heterogeneity in the  $I^2$ -measure. Therefore, this review has only reported the  $I^2$ -measure, instead of interpreting and labelling the  $I^2$ -measure by using cut-off points. Furthermore, in this review we limited the scope of outcomes to BMI, zBMI, waist circumference and weight status. We acknowledge the fact that the use of other anthropometric indicators (such as fat percentage) and more in-depth considerations of methods to assess obesity (such as the 95th percentile) may lead to additional insights in the future. Final limitations are that trial registries were not searched and that non-English studies were not included in this systematic review.

In line with recommendations by Brown and colleagues (2019) we performed this review to aggregate the current evidence on the long-term effects of school-based obesity prevention interventions.<sup>5</sup> We cross-checked the included RCTs in this review against those in the Cochrane review by Brown, and substantially expanded the evidence base by our more recent search and by allowing other designs than RCTs (6 studies). In comparison, from the 13 included RCTs in this review, 3 RCTs were also included in the Cochrane review, <sup>34,36,42</sup> for 5 RCTs we included studies with a longer follow-up of included studies in the Cochrane review, <sup>26,28,31,32,37</sup> and 5 RCTs were not included in the Cochrane review. <sup>27,35,38,43,44</sup>

Due to the limited number of available studies and the low quality of studies we cannot present clear evidence regarding the long-term effects of interventions on obesity-related outcomes. Based on the findings in this review, we have the following recommendations for future research. First of all, we emphasize the urgent need for higher quality follow-up studies to add to the knowledge base on the long-term effectiveness of primary school-based preventive interventions. Based on the shortcomings in quality of the reviewed studies, we recommend to prospectively register the studies, choosing appropriate analysis techniques, to adequately report on missing outcome data, to blind assessors and to adequately describe intervention and control group conditions. Clear information on control group content (attention control or usual practice) is important, because studies with attention

control help researchers to disentangle the beneficial effects of attention and interpersonal interactions from the intervention itself.<sup>57</sup> RCTs are often considered the golden standard for these effectiveness evaluation studies, 11 but long-term follow-up of RCTs might be challenging due to funding issues or because it is unethical to withhold children from effective interventions for more years. Furthermore, the longer the follow-up period, the more likely it will become that power or attrition issues arise that make it more difficult to reach sound evidence on effectiveness. Therefore, to assess the long-term impact of interventions, other types of study designs may give important insights as well. Natural experiments, quasi-experimental and observational studies might be considered more feasible due to logistic or practical issues or can be considered more ethical than randomizing children for years in a control group withholding them from future interventions which are possibly effective. 12,13 In addition, using data from existing registries or ongoing cohort studies might be a way to derive insights on the longterm effects of interventions beyond their intervention period. Furthermore, we recommend the use of predictive modelling combining intervention characteristics and individual participant data in future studies to more adequately determine the effect of different intervention components and in population subgroups. If researchers choose for RCT designs, they should take into account the consequences of participant attrition at follow-up for their statistical power. We recommend researchers to take this into account for their power calculations and sample size determination when setting up new intervention effect evaluation studies. When reviewing the included interventions in this review, it becomes clear that interventions with a duration of multiple school years are underrepresented. More studies into the long-term effectiveness of interventions with longer durations are therefore recommended. Furthermore, when more evidence from high quality studies becomes available in the future, it will be valuable to review in more detail the differences in intervention effectiveness between subgroups of the population (such as differences in socioeconomic status or weight status) in an updated review. Finally, we recommend researchers to conduct more long-term effect studies in which both obesity-related outcomes and changeable lifestyle factors are measured. These studies are valuable to gain more knowledge on the pathways that underlie behaviour change in youth. It is likely that the association between obesity prevention interventions and their long-term effect on obesity-related outcomes such as BMI, zBMI, WC and weight status is mediated by intermediate factors related to a healthy lifestyle. Caloric overconsumption, poor food choices, sedentary behaviour and low levels of physical activity are changeable factors for the prevention of childhood obesity.<sup>58-61</sup> A recent systematic review and meta-analysis by Nally and colleagues observed that primary school-based interventions have no clear effect on healthy lifestyle behaviours such as sedentary behaviour, moderate-to-vigorous physical activity and diet behaviour.<sup>62</sup> Therefore, future research is warranted to explore the actual change in lifestyle that precedes outcomes on overweight on the short and longer term in primary school-based interventions.

Based on our findings, we have the following recommendations for practice. Policymakers and intervention developers should bear in mind that evidence of sustained effects of many included interventions in this review is unclear. Besides methodological explanations, explanations can be found in the obesogenic environment that surrounds and shapes the health behaviour of children and adolescents, <sup>53,54</sup> opposing the accomplished effects immediately post-intervention. This may signify the need to implement prolonged interventions or additional interventions for children and adolescents to sustain and improve a healthy lifestyle. As a considerable number of the interventions showed immediate post-intervention effectiveness, more focus might also be needed on maintaining healthy behaviours. Whole system approaches with more focus on policies, systems and environments might be a promising way forward for policymakers in order to initiate and maintain behaviour among schoolchildren. <sup>63</sup> Therefore, a promising way for policy makers to intervene may therefore be policy, systems and environment changes supporting healthy behaviour in school and community settings.

## 5 | CONCLUSION

The results of this review contribute to knowledge on the long-term effects of obesity prevention interventions. It did not show clear evidence regarding the long-term effects of primary school-based obesity prevention interventions in children on obesity-related outcomes and certainty levels of the evidence were generally low. We recommend policy makers and intervention developers to implement prolonged interventions or additional interventions for children and adolescents to sustain a healthy lifestyle. Collaborating with researchers in early phases of intervention planning may be useful for this purpose Moreover, we emphasize the need for long-term effect evaluation studies using more innovative study designs. We specifically emphasize the urgent need for higher quality studies, to be able to reach stronger conclusions about the long-term effects of primary school-based obesity prevention interventions in children.

# **AUTHOR CONTRIBUTIONS**

Michel S. Smit, Mirte Boelens, Famke J. M. Mölenberg, Hein Raat and Wilma Jansen contributed to the design of the study and the development of the protocol for this study. Michel S. Smit and Mirte Boelens performed the selection process of articles, data extraction and analysed the data. Michel S. Smit mainly wrote the manuscript and Mirte Boelens, Famke J. M. Mölenberg, Hein Raat and Wilma Jansen contributed to the manuscript by critical revisions and giving comprehensive feedback on multiple drafts. Michel S. Smit, Mirte Boelens, Famke J. M. Mölenberg, Hein Raat and Wilma Jansen read and approved the final manuscript.

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#### SUPPORTING INFORMATION

Additional supporting information can be found online in the Supporting Information section at the end of this article.

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