SYSTEMATIC REVIEW



Economic Evaluations of Preventive Interventions for Dental Caries and Periodontitis: A Systematic Review

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Abstract

Objectives To critically examine the methods used for full economic evaluations of preventive interventions for dental caries and periodontitis.

Methods Published literature post-2000 was searched to April 2021. Based on a developed intervention classification framework for dental caries and periodontitis, only universal, selective or indicated interventions were included in this review. The Drummond 10-point checklist was used for quality appraisal.

Results Of 3,007 unique records screened for relevance, 73 studies were reviewed. Most model-based studies (61/73) used cost-effectiveness analysis (49%) or cost-benefit analysis (28%). Trial-based studies (16/73) commonly used cost-effectiveness analysis (59%). Four studies used both economic evaluation methods. Sixty-four papers (88%) were on dental caries, eight papers (11%) focused on periodontitis, and one paper (1%) included both oral diseases; 72% of model-based and 82% of trial-based studies were of good quality. The most frequently investigated dental caries preventive interventions were water fluoridation (universal intervention; cost-saving or cost-effective), fissure sealant and fluoride varnish (selective and indicated intervention; cost-effective) was the most frequently evaluated preventive intervention for periodontitis. Thirty percent of studies with a time horizon > 1 year did not apply an appropriate discount rate and 26% did not comprehensively discuss other important considerations beyond the technical analysis.

Conclusions Generic health outcome measures should be incorporated for economic evaluations on preventive interventions for dental caries and periodontitis, and an increased focus to prevent periodontitis using economic evaluation methods is needed to inform resource allocation and policy decision-making.

1 Introduction

The major contributors impacting quality of life due to oral disease are predominantly caused by advanced dental caries, periodontitis (gum disease) and advanced periodontitis. Advanced dental caries and advanced periodontitis affecting multiple teeth can lead to edentulism (total tooth loss). The 2017 Global Burden of Disease (GBD) study

estimated 2.3 billion people had dental caries in permanent (adult) teeth, 532 million children had dental caries in deciduous (baby) teeth, 796 million people have advanced periodontitis, and 267 million people have edentulism [1]. These statistics correspond to 1.6 million years lived with disability (YLD) for permanent teeth, 0.1 million YLD for deciduous teeth, 5.2 million YLD for severe periodontitis and 7.3 million YLD for edentulism [1].

Dental caries and periodontitis are experienced disproportionately by different populations in society [2–4], and share common risk factors with other noncommunicable diseases (NCDs) [5, 6]. Additionally, periodontitis can have a direct association with other NCDs including cardiovascular disease [7, 8], dementia and cognitive impairment [9, 10], diabetes [11, 12] and obesity [13, 14]. The 2017 GBD study reports that more economically developed countries had the lowest burden of dental caries and advanced periodontitis [1]. The burden of oral diseases

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Key Points for Decision Makers

Methods used for economic evaluations on preventive interventions for dental caries and periodontitis were diverse, including the cost components, the consequences, and health outcomes.

These differences make it difficult to enable comparability between interventions to inform health investment decisions.

Future research in the field should consider how to incorporate generic health outcome measures, increase focus on performing economic evaluations to prevent periodontitis, and consider health equity impacts.

is significant, yet oral health is often excluded from evidence-based public health policy [15-17].

Despite a large body of evidence demonstrating the clinical effectiveness of a broad range of preventive interventions for dental caries and periodontitis, few interventions are translated into routine clinical dental practice or public health programs [18–22]. Dentistry has remained treatment-dominated and a highly specialised area of healthcare, which, as a result, has not successfully tackled the global burden of oral diseases [5].

Evidence for the cost-effectiveness of interventions is also an important consideration to assist decision makers in determining investment decisions. In the past 10 years, seven systematic reviews of economic evaluations were focused on preventive interventions for dental caries and/or periodontitis [23–29]. Six systematic reviews were exclusively focused on dental caries, of which four reviews were on child populations [25–28]. One systematic review assessed interventions for dental caries, periodontitis and oral cancer [29].

When considering the focus of these reviews, five reviews were primarily concerned with investigating the quality of the economic evaluation studies [23–26, 29]. Four reviews provided an overall economic evaluation quality score using various checklists [23, 25, 27, 29], although determining study quality using a scoring system is not a best practice recommendation [30, 31]. One review did avoid reporting an overall score but developed and used a risk of bias grading assessment [24].

The most common preventive interventions that have been economically evaluated include water fluoridation [23, 29], fissure sealant applications [23, 26, 29], and the use of topical fluorides such as fluoride varnish applications and fluoride mouth rinses [24, 26, 28]. Only three out of the six systematic reviews that assessed these preventive interventions reported whether the preventive interventions were cost-effective [24-26]. This is due, in part, to the reviews' primary aim being to determine the quality of economic evaluation studies. In general, preventive interventions found to be cost-effective were water fluoridation, taxation of sugar sweetened beverages (SSBs), and anticipatory guidance inclusive of oral health education. Fissure sealant applications and interventions (with or without dental treatment) incorporating the use of topical fluorides (mouth rinses, gels and fluoride varnish applications) were shown to have mixed cost-effectiveness outcomes (either cost-effective or more costly and more effective). The lack of primary studies on economic evaluations for the prevention of periodontitis continues to be a gap in the literature. In addition, the economic evaluation methodology on prevention interventions for dental caries and periodontitis have not yet been fully explored. Therefore, the aim of this study was to conduct a systematic literature review to critically examine the methods used for full economic evaluations of preventive interventions for dental caries and periodontitis.

2 Methods

The study protocol was registered in May 2020 in PROS-PERO (CRD42020186409) and revised in October 2020 to ensure there was contemporary relevance.

2.1 Literature Search

Published literature was searched in May 2020 and updated in April 2021 for publications post-2000 using the following electronic databases: The CRD databases (DARE, NHS EED, HTA), EBSCO databases (CINAHL, ERIC, Global Health, OpenDissertations), EMBASE, Medline and Web of Science. Efforts were made to identify relevant unpublished 'grey' literature and conference proceedings through appropriate websites and databases such as Proquest Dissertations and Theses Global, OpenGrey, and EThOS databases for potentially relevant studies. Google Scholar and government websites were not searched due to the limitations of search replicability. Bibliographic information from identified relevant systematic reviews were examined for potentially applicable studies. Medical Subject Headings (MeSH) and free-text search terms were combined with key concepts relating to dental caries, periodontitis and economic evaluation. The adopted search terms and strategy are specified in Online Supplementary Material (OSM) Appendix 1.

2.2 Study Selection and Eligibility

The study eligibility criteria were based on the population, intervention, comparator and outcome (PICO) framework: P—all ages, I—Preventive interventions for dental caries and periodontitis, C—at least one strategy (the intervention or comparator) must be a preventive intervention or prevention is a primary focus (dental treatment can be a consequence), and O—dental caries or periodontitis and economic evaluation.

In this review, we adapted the intervention framework that has been commonly used in mental health prevention [32, 33] to develop the Intervention Classification Framework for Dental Caries and Periodontitis. The spectrum of interventions illustrated in Fig. 1 classifies six categories according to their descriptors as explained in OSM Appendix 2. Only universal, selective or indicated interventions were included in this review. Universal interventions target the whole population and not people identified based on individual risk for dental caries or periodontitis. Selective interventions target community population subgroups whose risk of developing dental caries or periodontitis is higher than average. Indicated interventions target high-risk individuals or those with initial stages of dental caries or periodontitis.

Other inclusion parameters included full-texts, English language publication, model-based or trial-based full economic evaluations (primary or secondary data), original empirical research, peer-reviewed and non-peer-reviewed publications (e.g., theses, government reports). Full economic evaluation is a systematic framework that allows the incremental analysis of alternative interventions in terms of both costs (resource use) and consequences (outcomes, effects) [34].

Other specific exclusion criteria include:

- Studies published earlier than 2000.
- Partial economic evaluations, where the incremental analysis of alternative interventions was not performed.
- Studies that made comparisons of different dental materials or clinical techniques, investigated the different use of the dental workforce or payment schemes.
- Studies that did not have a primary focus on preventive interventions.



Fig. 1 Interventions classification framework for dental caries and periodontitis, adapted from Mrazek and Haggerty [32]

• Literature/systematic reviews, letters, editorials, conference abstracts and commentaries.

The screening process and quality appraisal were evaluated for their level of agreement according to Cohen's Kappa values whereby a score < 0.40 corresponds to poor, 0.40–0.74 is considered fair to good, and 0.75–1 is defined as perfect agreement [35]. It was agreed that studies pre-2000 would be excluded considering that economic evaluations in oral health research have considerably increased from the year 2000 onwards [36]. Potential papers identified using the literature search strategy were imported and duplicate records removed using Endnote X9.2 (Clarivate Analytics). The remaining papers for screening were imported and managed using Covidence. Two reviewers (TMN and UT) independently screened title/abstract and full-text screening. Any differences in agreement were discussed and a consensus reached.

2.3 Data Extraction

TMN completed the data extraction for studies into Covidence using the following parameters:

- Publication year, author, classification of preventive intervention, country context, data source (primary/secondary data), economic evaluation study type (model/ trial/combination)
- B. Type of economic evaluation (e.g., cost-benefit analysis (CBA), cost-consequence analysis (CCA), cost-minimisation analysis (CMA), cost-effectiveness analysis (CEA), cost-utility analysis (CUA) or combination and return on investment (ROI) studies.
- C. Economic evaluation characteristics including perspective, reference year, time horizon, discount rate for costs and outcomes, cost categories.
- D. Intervention and comparator description.
- E. Population characteristics including age of target population, and number of participants.
- F. Reported outcomes including:
 - Dental outcomes (e.g., caries free, decayed teeth surfaces (ds/DS), decayed teeth (dt/DT), decayed, missing, and filled surfaces (dmfs/DFMS), decayed, missing, and filled teeth (dmft/DMFT), proportion of bleeding sites, retained-tooth per year, qualityadjusted tooth years (QATYs),
 - Health outcomes (e.g., disability-adjusted life years (DALYs) and quality-adjusted life years (QALYs), and
 - Economic evaluation outcomes (e.g., incremental cost-effectiveness ratio (ICER), cost-benefit ratio (CBR), and ROI.

2.4 Cost-Effectiveness

Value for money judgments were made based on whether the preventive intervention was cost-saving, dominant, or was below a willingness-to-pay (WTP) threshold. For CEA studies that calculated the ICER, where the intervention was more costly and more effective, cost-effectiveness was 'not stated' due to the absence of WTP thresholds for dental outcomes. If a WTP threshold was used to judge costeffectiveness for CUA studies, as these can differ between countries, this was included in the review.

2.5 Quality Appraisal

The methodological quality of eligible model-based and trial-based studies was appraised using the Drummond 10-point checklist, including the 32 sub-questions, which informed the response of the overarching question (answer options being 'yes', 'no' or 'unsure') [34]. The Drummond 10-point checklist has previously been applied for systematic reviews of economic evaluations on oral health interventions [24, 27, 36, 37]. Quality appraisal was also used to assist in identifying the methodological characteristics of the economic evaluations included in this review. All studies were appraised by one reviewer (TMN), and a second appraisal was provided by two other independent reviewers (50:50; UT and LL). Any differences in the quality appraisal scoring were discussed between the reviewers to reach a consensus, with unresolved issues mediated by the third reviewer. Individual study and overall study quality according to the preventive interventions were classified using Method 1 from the Gonzalez-Perez (2002) criteria [38] for the Drummond 10-point checklist [34]: low quality (0–0.69), moderate quality (0.7-0.89) and good quality (0.9-1.0).

3 Results

The literature search process is summarised using the PRISMA guidelines (Fig. 2) [39]. Of the 5,133 records identified, the duplicate study removal process resulted in 3,007 unique records, with 2,685 records excluded at the title/abstract screening stage (TMN/UT; Cohen's Kappa: 0.605). Four records were identified via bibliographic citation search. Of the remaining 322 records subjected to full-text screening (TMN/UT; Cohen's kappa: 0.430), 29 records were automatically removed since they reported abstracts only or the full-texts could not be retrieved. The final number of studies meeting the inclusion eligibility criteria was 73 full-text articles. Both title/abstract and full-text screening had fair to good agreement between reviewers.

3.1 Findings Summary

A summary of the study characteristics is reported in Table 1, with the individual details presented in OSM Appendix 3. Based on the full-text analysis, 64 studies (88%) were related to dental caries, eight (11%) to periodontitis, and one study (1%) investigated impacts on both dental caries and periodontitis. Most studies (90%) were in high-income countries including the USA (n = 17), Australia (n = 15), and the UK (n = 11). Of the 73 full-text papers, there were 34 categories of preventive interventions for dental caries and/or periodontitis; ten were universal interventions, nine were selective interventions, and 15 were indicated interventions (Table 2).

The most frequently economically evaluated preventive interventions for dental caries were water fluoridation (universal), fissure sealant and fluoride varnish (selective and indicated). For periodontitis, the most common preventive intervention economically evaluated was supportive periodontal therapy with oral health education. The target population for these interventions were mainly only for children (64%) followed by both adults and children (22%).

A summary of the characteristics of included studies is reported in Table 3. The most common economic evaluation approaches used were CEA and CBA (47% and 25%, respectively). Model-based studies were the most common approach for economic evaluation (78%), with a few studies using both model-based and trial-based studies (5%).

Almost half (49%) of model-based studies used the societal perspective and 44% used the healthcare perspective, compared with 35% for trial-based studies that used either the healthcare or the societal perspective. A greater proportion of trial-based studies (29%) compared with model-based studies (5%) performed economic evaluation from both the healthcare and societal perspectives. The most common cost categories were the cost of direct restorations only (33%) and direct treatment costs inclusive of restorations, extractions and other direct costs for dental caries (36%) for modelbased studies, where the latter cost category was the most used for trial-based studies. Other direct costs for dental caries include the cost of root canal treatments, crowns, bridges and implant crowns, noting that these dental treatment procedures may be caused by trauma, cracked tooth syndrome, and severe stages of periodontitis rather than because of dental caries.

The most common dental outcome measures for dental caries were dmfs/DMFS, dmft/DMFT averted, ds/DS averted and caries-free proportion of the population, months or persons. The most common dental outcomes for periodontitis were retained-tooth per year and the proportion of teeth with bleeding sites. CUA model-based and trial-based studies (14%) used either DALYs for dental caries and QALYs for dental caries and periodontitis as the health outcome



Fig. 2 Results of the systematic review search strategy using the PRISMA guidelines [39]

measures. One study quantitatively used a health equity outcome measure, which evaluated whether the intervention increased access to dental services for children from lowincome households [94].

3.2 Derivation of Disability and Utility Weights for General Health Outcome Measures

The use of disability and utility weights in the selected studies to quantify DALY averted and QALY gained was variable. Two studies measured DALY using the disability weight of 0.057 for symptomatic dental caries according to the 2003 Australian Burden of Disease study. It includes two time periods spent with disability, an initial phase where the individual experiences intermittent pain and the terminal phase where the individual experiences constant pain. This approach has later been adopted by the 2017 GBD study [1]. One study used the disability weight of 0.012 for dental caries derived from the 2010 GBD study [51] but did not state if the initial and terminal phase experience of pain was considered.

Except for two studies [52, 53], there were generally two approaches to quantify QALYs: (1) one minus the

disability weight used in GBD studies or (2) measured using quality-of-life survey instruments for trial-based studies. One study used the disutility weight of 0.720 for otitis media (middle ear infection) [54], and one study used the disutility weight for different severity of dental caries experience (dmft/DMFT) [55], derived from the New Zealand Burden of Disease study [56]. For trial-based studies, one study used the Child Health Utility Nine-Dimension (CHU-9D) instrument, either completed by the parent/ primary caregiver as a proxy measure [57] or via the children directly [40], and another study used the EQ-5D-3L instrument to derive QALY utility weights for adults [58]. One study used 'expert opinion' for the utility weights for periodontitis, and another study represented QALY as the aggregate preference tooth utility weight of eight permanent molar teeth to represent the individual [52].

3.3 Willingness-To-Pay Thresholds

Various hypothetical WTP thresholds for dental outcomes were considered by seven studies [48–50, 59–62]. Two Chilean studies have suggested a WTP threshold of 20,000 Chilean pesos per QATY gained or caries-free person, based on the cost of a restoration from the healthcare

Table 1 A summary of the study characteristics included in the systematic review

Summary	Characteristic	No. stuc N = (%)	. of dies = 73
Country context*	High income		
	Australia [52, 57, 63-65, 67, 68, 70, 87, 92-94, 99, 103, 108]	15	(19%)
	Canada [96, 100]	2	(3%)
	Chile [44, 59, 61, 82, 85]	5	(6%)
	Finland [60]	1	(1%)
	Germany [49, 50, 62, 89, 90, 108, 110]	7	(9%)
	Ireland [71, 108]	2	(3%)
	Japan [97, 108]	2	(3%)
	Netherlands [86, 105]	2	(3%)
	New Zealand [55, 73, 79]	3	(4%)
	Norway [109]	1	(1%)
	Spain [108]	1	(1%)
	Sweden [48, 106, 107]	3	(4%)
	UK [40, 54, 58, 84, 95, 102, 104, 108, 111–113]	12	(15%)
	USA [41, 43, 45–47, 53, 69, 74, 76, 88, 91, 98, 101, 102, 108, 123, 124]	17	(21%)
	Subtotal	72	(90%)
	Upper-middle income		
	Brazil [42]	1	(1%)
	Iran [72]	1	(1%)
	Peru [80]	1	(1%)
	South Africa [75, 78]	2	(3%
	Thailand [81]	1	(1%)
	Subtotal	6	(8%)
	Low-middle income		
	Sri Lanka [108]	1	(1%)
	Low income		
	Nepal [83]	1	(1%)
Target oral disease	Dental caries [40-50, 52, 54, 55, 57-65, 67-69, 71-91, 93-101, 103-107, 111, 112, 123, 124]	65	(88%)
	Periodontitis [53, 92, 102, 108–110, 113]	7	(11%)
	Caries and periodontitis [70]	1	(1%)
Target population	Adults [50, 53, 58, 64, 88, 92, 102, 108–110]	10	(14%)
	Children^ [40-49, 52, 54, 57, 59-62, 67-69, 71, 74, 80-85, 90, 91, 93-101, 103-107, 112, 123, 124]	47	(64%)
	Adults + children^ [55, 63, 65, 70, 72, 73, 75–79, 86, 87, 89, 111, 113]	16	(22%)
Economic evalua- tion study type	Model-based [41, 43-47, 49, 50, 52-55, 57, 59, 61-63, 67-91, 93, 95, 96, 98-101, 103, 105, 108-111, 123, 124]	57	(78%)
	Trial-based [48, 58, 60, 92, 94, 97, 102, 104, 106, 107, 112, 113]	12	(16%)
	Model + trial [40, 42, 64, 65]	4	(5%)
Data source	Primary [40, 42, 48, 58, 60, 63, 64, 92, 94, 97, 102, 104–107, 112, 113]	17	(23%)
	Secondary [41, 43-47, 49, 50, 52-55, 57, 59, 61, 62, 65, 67-91, 93, 95, 96, 98-101, 103, 108-111, 123, 124]	58	(77%)

N total number of studies

*One study reported economic evaluations for eight different country contexts [108]

^Children were defined as < 18 years old

perspective [59, 61], and three studies have referenced AU\$3,300 per restoration averted [63–65], derived from a study estimating the average lifetime average costs of probable avoided restoration per person [66]. Two other

studies used various WTP thresholds using proposed dental outcomes [48, 60]. The WTP threshold of AU50,000 per DALY averted [67, 68], £20,000 per QALY gained [40, 58], and the 2014 US gross domestic gross product

Table 2	The categories of	preventive interventions th	at were economicall	y evaluated for dental	caries and periodontitis

Preventive intervention description		nber of nomic uations 100 (%)	Target oral disease	Economic evaluation outcome	Overall quality of studies*
Universal interventions $(N = 28)$					
20% sugar-sweetened beverages tax vs. no intervention [86, 87]	2	(2%)	Dental caries	Cost-effective or Cost-saving	Good
Bacterial screening saliva testing vs. standard care [91]	1	(1%)	Dental caries	N/S	Moderate
Ban on sugar-sweetened beverages sales vs. no interven- tion [88]	1	(1%)	Dental caries	Cost-saving	Good
Fluoridation of toothpastes vs. no intervention [83]	1	(1%)	Dental caries	Cost-saving	Moderate
Front-of-package food labelling vs. no intervention [89]	1	(1%)	Dental caries	Cost-effective	Good
Increasing the use of chewing gum vs. no intervention [90]	1	(1%)	Dental caries	Cost-saving	Good
Interleukin-1 genetic testing vs. standard care [53]	1	(1%)	Periodontitis	N/S	Good
Milk fluoridation vs. no intervention [81, 82, 85]	3	(2%)	Dental caries	Cost-effective	Good
Salt fluoridation vs. no intervention [80, 85]	2	(2%)	Dental caries	Cost-effective	Good
Water fluoridation vs. no intervention [55, 67, 68, 70–79, 84, 85]	15	(12%)	Dental caries and periodontitis [#]	Cost-effective or Cost-saving	Moderate
Selective interventions $(N = 29)$					
Targeted postage of oral hygiene products vs. no interven- tion [95]	1	(3%)	Dental caries	N/S	Low
Targeted pre/primary school-based dental check-up vs. standard care [94]	1	(3%)	Dental caries	Cost-effective	Good
Targeted pre/primary school-based fluoride varnish by dental/non-dental primary care providers with/without dental screening [44, 54, 61, 96]			Dental caries		
vs. counselling [44]	2	(7%)		Not cost-effective or N/S	Good
vs. no intervention [54, 61]	2	(7%)		Cost-effective or N/S	Moderate
vs. standard care [96]	1	(3%)		Cost-effective	Moderate
Targeted primary school-based fissure sealant program with/without fluoride mouthrinse (intervention)			Dental caries		
vs. no intervention [59, 84, 85, 97, 98]	5	(17%)		Cost-effective or Cost-saving or N/S	Moderate
vs. standard care [69, 99, 100]	4	(14%)		Cost-effective or N/S	Good
vs. targeted school-based toothbrushing program [42]	1	(3%)		N/S	Good
Targeted primary school-based fissure sealant program (comparator)			Dental caries		
All seal vs. risk-based [98, 101]	2	(7%)		N/S	Moderate
No intervention vs. risk-based [98, 101]	2	(7%)		Not cost-effective	Moderate
Targeted primary school-based fluoride gel vs. no inter- vention [85]	1	(3%)	Dental caries	N/S	Good
Targeted primary school-based fluoride mouth rinse vs. no intervention [85]	1	(3%)	Dental caries	Cost-effective	Good
Targeted supervised toothbrushing program vs. no inter- vention [42, 54, 85]	2	(7%)	Dental caries	N/S	Moderate
Targeted telehealth or home visit oral health education vs. standard care [57, 84, 93]	3	(10%)	Dental caries	Cost-saving or Cost-effective	Moderate
Targeted community-based group oral health education vs. chairside oral health education [92]	1	(3%)	Periodontitis	Cost-saving	Good
Indicated interventions $(N = 42)$					
Artificial intelligence for intra-oral radiographic detection of dental caries vs. no intervention [49]	1	(2%)	Dental caries	Cost-effective	Good
Oral prophylaxis and oral health education vs. fissure sealant and fluoride varnish [103]	1	(2%)	Dental caries	Not cost-effective	Moderate

Table 2 (continued)

Preventive intervention description		nber of nomic luations : 100 (%)	Target oral disease	Economic evaluation outcome	Overall quality of studies*	
Fluoride and xylitol supplement, oral prophylaxis and fluoride/chlorhexidine varnish vs. standard care [60, 107]	2	(5%)	Dental caries	N/S	Good	
Fissure sealant vs. no intervention [41, 45–47]	5	(12%)	Dental caries	N/S	Moderate	
Fluoride varnish vs. fissure sealant [40, 41]	2	(5%)	Dental caries	Cost-effective or Not cost-effective	Moderate	
Fluoride varnish vs. standard care [48, 52, 62, 105, 106, 112]	6	(14%)	Dental caries	N/S	Good	
Minimally invasive dentistry (intervention)			Dental caries			
vs. no intervention [104]	1	(2%)		N/S	Good	
vs. standard care [63-65, 104, 105]	5	(12%)		Not cost-effective or N/S	Good	
Standard care vs. minimally invasive dentistry [104]	1	(2%)	Dental caries	Not cost-effective	Good	
Slow releasing fluoride glass devices program [84]	1	(2%)	Dental caries	Cost-saving	Low	
Resin infiltration vs. fluoride varnish and/or oral health education [50]	1	(2%)	Dental caries	N/S	Moderate	
Varied intervals of dental check-ups e.g., 3-, 9-, 12-, 24-, 36-monthly, risk-based			Dental caries			
vs. standard care [58, 111]	7	(17%)		N/S	Good	
24-monthly vs. risk-based [58]	1	(2%)		Cost-effective	Good	
Regular supportive periodontal therapy			Periodontitis			
vs. irregular supportive periodontal therapy [109, 110]	2	(5%)		N/S	Moderate	
vs. tooth removal and implant crown replacement [110]	1	(2%)		N/S	Good	
Supportive periodontal therapy by dental specialist vs. general dental practitioner [108]	1	(2%)	Periodontitis	N/S	Good	
Varied intervals of supportive periodontal therapy and standard oral health education (e.g., 12-monthly)			Periodontitis			
vs. standard care [102]	1	(2%)		Cost-saving	Good	
Varied intervals of supportive periodontal therapy and tai- lored oral health education (e.g., 6-monthly, 12-monthly, risk-based)			Periodontitis			
6-monthly vs. standard care [102]	1	(2%)		N/S	Good	
12-monthly vs. standard care [102]	1	(2%)		N/S	Good	
Risk-based vs. standard care [102]	1	(2%)		N/S	Good	

N = total number of economic evaluations, N/S not stated, Standard care refers to treatment as usual with 6-monthly dental check-ups

[#]Only one study on water fluoridation economically evaluated dental caries and periodontitis [70]

*Overall quality of studies using the Gonzalez-Perez (2002) criteria [38] for the Drummond 10-point checklist [34]

per capital (US\$54,639 per DALY averted) [69] were used for CUA studies in Australia, the UK and the USA, respectively.

3.4 Universal Interventions

The majority of universal interventions were reported to be cost-effective or cost-saving. These universal interventions included various forms of topical fluorides [55, 67, 68, 70–85], interventions targeting the reduction of sugar consumption such as SSBs taxation or ban on selling SSBs at the workplace [86–89], increasing use of chewing sugar-free gum [90], and bacterial screening saliva test for virulent levels of mutans streptococci [91]. For interleukin-1 genetic screening to prevent periodontitis for persons aged 35 years, the ICER was calculated but the cost-effectiveness was not stated [53].

3.5 Selective Interventions

For selective interventions, targeted community-based group oral health education for adults [92], targeted primary schoolbased fluoride mouth-rinse program [85], targeted telehealth or home visit oral health education for young children [57,

Table 3 A summary of characteristics for model-based and trial-based economic evaluation studies

Characteristics	Mod Ecor N =	odel-based onomic evaluations = 61* (%)		
Economic evaluation approach				
CBA [55, 70, 72, 74–78, 83, 84, 86–88, 90, 91, 123, 124]	17	(28%)		
CEA [41-47, 49, 59, 61-65, 73, 79-82, 85, 93, 95, 96, 98, 100, 101, 105, 108, 109]	30	(49%)		
CUA [53, 54, 57, 67–69]	6	(10%)		
CBA and CEA [50, 71, 89, 99, 111]	5	(8%)		
CEA and CUA [40, 52]	2	(3%)		
Perspective				
Healthcare [41, 43, 45–47, 49, 50, 52–54, 59, 61–65, 84, 85, 87, 90, 91, 101, 109–111, 123, 124]	27	(44%)		
Societal [42, 44, 55, 57, 67-83, 86, 89, 93, 95, 96, 98-100, 108]	30	(49%)		
Healthcare and societal [40, 88, 105]	3	(5%)		
Reference year				
N/R [54, 75, 84, 91, 95, 96, 123]	7	(11%)		
< 2000 [53, 74, 78, 79, 82, 98, 99]	7	(11%)		
2000–2005 [43, 45, 46, 67, 68, 70, 77, 81, 83, 101, 111]	11	(18%)		
2006–2010 [64, 65, 80, 85, 100, 108]	6	(10%)		
2011–2015 [40–42, 44, 47, 49, 50, 52, 55, 57, 63, 69, 72, 73, 76, 87, 90, 93, 105, 109, 110]	21	(34%)		
2016–2020 [59, 61, 62, 71, 86, 88, 89, 124]	8	(13%)		
Time horizon				
N/R [73]	1	(2%)		
< 5 years [125]	14	(23%)		
5-10 years [40, 41, 45, 47, 57, 59, 63, 75, 80, 81, 83, 85, 87, 89, 93, 96, 98-101, 123]	21	(34%)		
> 10 years [49, 50, 52, 53, 55, 62, 67, 68, 70, 71, 74, 76, 77, 79, 86, 90, 108–110]	19	(31%)		
Mixed (< 5 and 5–10 years) [124]	1	(2%)		
Mixed (< 5 and > 10 years) [63, 64]	2	(3%)		
Mixed $(5-10 \text{ and } > 10 \text{ years})$ [88, 111]	2	(3%)		
Cost categories—healthcare and societal perspective [#]				
Direct treatment costs—N/R [41, 44, 95]	3	(5%)		
Direct treatment costs—restorations only [42, 61, 67–69, 72–75, 78, 79, 83, 87, 89, 98–101, 105, 111]	20	(33%)		
Direct treatment costs—restorations and extractions only [40, 47, 59, 71, 76, 80–82, 85]	9	(15%)		
Direct treatment costs—restorations, extractions and other direct costs for dental caries [43, 45, 46, 49, 50, 52, 54, 55, 57, 62–65, 77, 84, 86, 90, 91, 93, 96, 123, 124]	22	(36%)		
Direct treatment costs—restorations, extractions and other direct costs for dental caries and periodontitis [70, 88, 110]	3	(5%)		
Direct treatment costs for periodontitis only [53, 108, 109]	3	(5%)		
Cost categories—societal perspective				
Direct societal costs—N/R [87]	1	(2%)		
Direct societal costs (intervention only) [42, 55, 67, 68, 71, 72, 75, 76, 78, 79, 83, 86, 89, 95, 96, 99]	15	(25%)		
Direct and indirect societal costs (productivity costs lost, travel costs, etc.) [40, 44, 57, 69, 70, 73, 74, 77, 80–82, 85, 88, 93, 98, 100, 105, 108]	17	(28%)		
Dental outcomes measures				
Decayed teeth surfaces/teeth averted (ds/DS) ^[41, 42, 50, 57, 70, 76–79, 89, 91, 93, 96, 98]	13	(21%)		
Decayed, missing and filled teeth surfaces/teeth averted (dmfs/DMFT, dmft/DMFT) [52, 55, 62, 64, 65, 71–75, 80–85, 87, 90, 95, 99, 105, 111, 124]	23	(38%)		
Caries free (%/months/person) [40, 43, 44, 61, 86, 95, 100, 101]	8	(13%)		
Clinical attachment loss averted [108]	1	(2%)		
QATY [40, 46, 59]	3	(5%)		
Restorative/extraction/episode of pain/hospital/event averted [43, 47, 63, 95, 123]	5	(8%)		
Retained-tooth per year [49, 62, 109, 110]	4	(7%)		
Tooth loss averted [108]	1	(2%)		

Table 3 (continued)				
Characteristics	Moo Eco N =	Model-based Economic evaluations $N = 61^*$ (%)		
Generic health outcome measures				
DALY [67-69]	3	(5%)		
QALY [40, 52–55, 57, 88]	7	(11%)		
Characteristics	Tri ecc N =	al-based pnomic evaluations = 17* (%)		
Economic evaluation approach				
CBA [102]	1	(6%		
CEA [42, 48, 60, 64, 65, 104, 106, 107, 112, 113]	10	(59%)		
CMA [92, 103]	2	(12%)		
CBA and CEA [97]	1	(6%)		
CBA and CUA [58]	1	(6%)		
CEA and CUA [40, 94]	2	(12%)		
Perspective				
Healthcare [60, 64, 65, 92, 103, 104]	6	(35%)		
Societal [40, 58, 102, 106, 112, 113]	6	(35%)		
Healthcare and Societal [42, 48, 94, 97, 107]	5	(29%)		
Reference year	U	(_>,\v)		
< 2000 [103, 107]	2	(12%)		
2000–2005 [60, 97]	2	(12%)		
2006–2010 [64, 65, 92, 113]	4	(24%)		
2011-2015 [40, 42, 94, 102, 106, 112]	6	(35%)		
2016-2020 [48, 58, 104]	3	(18%)		
Time horizon	5	(10/0)		
< 5 years [40] 42 48 58 60 64 65 92 94 102–104 106 112 113]	15	(88%)		
5_10 vers [107]	1	(6%)		
Mixed (< 5 and 5-10 years) [97]	1	(6%)		
Cost catagories healthcare and societal nerspective	1	(070)		
Direct treatment costs N/P [103, 106]	2	(12%)		
Direct treatment costs - rectorations only [42, 48, 107]	2	(12%)		
Direct treatment costs—restorations and extractions only [40]	1	(10%) (6%)		
Direct treatment costs—restorations and extractions only [40]	1	(0%)		
Direct treatment costs—restorations, extractions and other direct costs for dental carles [00, 04, 03, 94, 97, 104, 112]	/	(41%)		
Direct treatment costs for periodontitis only [02, 102, 112]	1	(0%)		
Cost catagorias sociatal perspective	5	(10%)		
Direct societal costs (intervention only) [42, 48, 97]	3	(18%)		
Direct and indirect societal costs (productivity costs lost travel costs atc.) [40, 58, 94, 102, 106, 107, 112, 113]	8	(13%)		
Dental outcomes measures	0	(4770)		
Decrya teeth surfaces/teeth overted $(ds/DS)^{1/2}$ 103 112]	3	(18%)		
Decayed missing and filled teeth surfaces/teeth averted	2	(10%)		
(dmfs/DMFT, dmft/DMFT) [48, 60, 64, 65, 94, 97, 106, 107]	0	(4770)		
Caries free (%/months/person) [40, 112]	2	(12%)		
Bleeding sites (%) [92, 102]	2	(12%)		
QATY [40, 94]	2	(12%)		
Restorative/extraction/episode of pain/hospital/event averted [104, 112]	2	(12%)		
Successful case managed for periodontitis [113]	1	(6%)		

Generic health outcome measures

Table 3 (continued)	
Characteristics	Trial-based economic evaluations N = 17* (%)
QALY [40]	1 (12%)
Other outcome measures	
Per 1% card-holder reached [94]	1 (6%)

N total number of studies/interventions, *N/R* not reported, *CBA* cost-benefit analysis, *CMA* cost-minimisation analysis, *CEA* cost-effectiveness analysis, *CUA* cost-utility analysis, *dmfs/DMFS*⁺ number of decayed, missing, and filled teeth surfaces for the deciduous or permanent dentition, *dmft/DMFT*⁺ number of decayed, missing, and filled teeth for the deciduous or permanent dentition, *ds/DS*⁺ number of decayed teeth surfaces for the deciduous or permanent dentition, *dt/DT*⁺ number of decayed teeth for the deciduous or permanent dentition, ⁺ lower-case letters refer to the deciduous dentition and upper-case letters refer to the permanent dentition, *DALY* disability-adjusted life years, *QATY* quality-adjusted tooth years, *QALY* quality-adjusted life years

*Four studies performed model-based and trial-based economic evaluations [40, 42, 64, 65]

[#]Four studies did not report the intervention costs [54, 87, 88, 90]

'Includes dental outcomes exclusively for occlusal or proximal surfaces

84, 93], and targeted pre/primary school-based dental checkup programs [94], were cost-saving or cost-effective.

The cost-effectiveness was not stated for targeted postage of oral hygiene products [95], targeted primary school-based fluoride gel [85], and targeted supervised toothbrushing program [42, 54, 85]. There were mixed cost-effectiveness results for targeted pre/primary school-based fluoride varnish by dental/non-dental primary care providers with/without dental screening [44, 54, 61, 96] and targeted primary school-based fissure sealant program with/without fluoride mouth rinse [42, 59, 69, 84, 85, 97–101]. In the context of targeted primary school-based fissure sealant programs, no intervention versus risk-based (fissure sealant application was based on the patient's dental caries risk) interventions was not cost-effective [98, 101].

Table 4 The quality appraisal results using the Drummond 10-point checklist [34]

Drummond 10-point checklist question		Publication year								
	2000-2010 N = 24			2011-2021 N = 49			2000-2021 N = 73			
	Yes	No	Can't tell	Yes	No	Can't tell	Yes	No	Can't tell	
Q1. Was a well-defined question posed in answerable form?	92%	4%	0%	98%	2%	0%	96%	3%	1%	
Q2. Was a comprehensive description of the competing alternatives given? (i.e., can you tell who did what to whom, where and how often?)	92%	4%	4%	100%	0%	0%	97%	1%	1%	
Q3. Was the effectiveness of the programmes or services established?	96%	0%	4%	100%	0%	0%	99%	0%	1%	
Q4. Were all the important and relevant costs and consequences for each alternative identified?	83%	0%	13%	94%	0%	6%	90%	0%	10%	
Q5. Were costs and consequences measured accurately in appropriate physical units prior to valuation (e.g., hours of nursing time, number of physician visits, lost work days, gained life-years)?	96%	0%	4%	96%	2%	2%	96%	1%	3%	
Q6. Were costs and consequences valued credibly?	96%	0%	0%	86%	4%	10%	89%	3%	7%	
Q7. Were costs and consequences adjusted for differential timing?	67%	25%	8%	71%	20%	8%	70%	22%	8%	
Q8. Was an incremental analysis of costs and consequences of alterna- tives performed?	92%	4%	0%	96%	4%	0%	95%	5%	0%	
Q9. Was uncertainty in the estimates of costs and consequences adequately characterised?	79%	21%	0%	90%	10%	0%	86%	14%	0%	
Q10. Did the presentation and discussion of study results include all issues of concern to users?	58%	38%	4%	82%	16%	2%	74%	23%	3%	

N number of studies

3.6 Indicated Interventions

Under indicated interventions, the use of artificial intelligence for the detection of dental caries on dental intra-oral radiographs was cost-effective [49]. Supportive periodontal therapy and standard oral health education provided 12-monthly versus 6-monthly [102], and the use of slowreleasing fluoride glass devices (glass ionomer cements) for children at high risk for dental caries risk [84] were costsaving. The economic evaluation outcomes were variable for the remaining studies, either cost-effective, not costeffective or not stated [40, 41, 45–48, 50, 52, 58, 62–65, 92, 98, 100–113]. Oral prophylaxis and oral hygiene education versus fissure sealant and fluoride varnish application was not cost-saving according to the CMA [103], and standard care versus minimally invasive dentistry was not cost-effective [104].

3.7 Quality Appraisal

The results of the quality appraisal using the Drummond 10-point checklist are reported in Table 4 regarding the sub-questions and Cohen's Kappa values. The study quality according to model-based and trial-based studies is shown in Table 5. Generally, the study quality for economic evaluations improved for studies published after 2010. For all 73 studies, some criteria of the Drummond 10-point checklist were not met, including 30% of studies not applying an appropriate discount rate (question 7) when the time horizon is greater than 1 year and 26% of the studies did not comprehensively discuss other important considerations beyond the technical analysis (question 10). Most model-based (72%) and trial-based studies (82%) were of good quality. In terms of quality appraisal assessment consistency, there was poor inter-rater agreement using the Drummond 10-point checklist (TMN/UT Overall Cohen's kappa: 0.164; TMN/LL Overall Cohen's kappa: 0.148; individual question Cohen's Kappa scores is reported in OSM Appendix 4).

Table 5 A summary of the study quality using the Gonzalez-Perez[38] criteria for the Drummond 10-point checklist for model-basedand trial-based studies[34]

Study quality	Model- $N = 61$	based studies	Trial-based studies $N = 17$			
Good	44	(72%)	14	(82%)		
Moderate	14	(23%)	3	(18%)		
Poor	3	(5%)	-			

N number of studies

4 Discussion

This study found that there was a significant number of peerreviewed publications on the economic evaluation of preventive interventions for dental caries, but there were limited publications on economic evaluation for periodontitis. Most of the studies (90%) were conducted in high-income countries. The choice in the selection of parameters used for economic evaluation was variable throughout the studies, including the time horizon, cost categories, and the chosen dental and health outcome measure(s). In general, universal interventions such as water fluoridation were cost-effective and cost-saving. Selective interventions were either costeffective or more costly and more effective, and indicated interventions had mixed cost-effectiveness.

Generally, the studies included in the review were demonstrated to be of moderate or good quality. There was generally poor inter-rater reliability using the Drummond 10-point checklist, but this is likely to be influenced by the assessor's interpretation of the question rather than the quality appraisal instrument itself [35]. Two of the three assessors had previous experience using the Drummond 10-point checklist, and two of three assessors have a background in dentistry. The poor level of inter-rater agreement highlights the importance of using at least two assessors, which may have influenced the quality appraisal of each study in either direction.

Our updated review noted there was a diversity of dental and generic health outcomes used in the studies included in this review and identified that there was sparse evidence for cost-effective interventions to prevent periodontitis [29, 114–116], which is a consistent finding from previous reviews [23–29]. Different outcome measures do not allow comparison between interventions or diseases to guide resource allocation. Our review found very few economic evaluations were conducted in low- and low-middle-income countries. Although quantitative outcome measures that capture health equity were not specifically investigated in our review, we found one study [94, 117] that incorporated it in their analysis. The issue of health equity is missing from previous reviews.

4.1 Interpretation and Implication of Outcome Measures

The use of dental outcome measures for economic evaluation has considerable limitations. For example, some authors compared preventive interventions using dental outcomes exclusively for occlusal or proximal surfaces [41, 48, 50, 98, 103], and described them as cavities [42, 96] or caries lesions [49, 57, 86, 89]. There were many studies that used the term caries increment with dental outcomes reported as either ds/DS, dt/DT, dmfs/DMFS and dmft/DMFT [43, 44, 48, 62, 64, 68, 71, 74–76, 85, 86, 89, 98, 105–107]. The nuance of terminology in dentistry to describe dental caries as numbers of tooth surfaces or teeth is distinct, as opposed to dental caries incidence in epidemiology, where an individual develops at least one tooth with dental caries. These measurement differences have implications for intervention comparability. With regards to periodontitis, there were variations in using tooth-specific or individual person measures such as retained-tooth per year, percentage of bleeding sites, and successful case managed.

In addition, the dmfs/DMFS and dmft/DMFT are only composite measures for dental caries since decayed teeth (d/D) is only one component of these indices [118]. Despite the limitations of both indices, the dmft/DMFT index is incorporated in the methodology used by the 2017 Global Burden of Disease study to calculate dental caries incidence given it is the most widely used instrument to measure for dental caries experience [1]. In brief, based on longitudinal studies, the population incremental difference in the dmft/ DMFT index for the initial and subsequent dental check-up is accepted as the number of dental caries incidence. These differences in outcomes are a limitation in quantifying the true measure of dental caries incidence. Adding to the complexity, many studies using CEA approaches could not be assessed for cost-effectiveness due to an absent WTP threshold using dental outcomes. This makes it difficult to interpret the cost effectiveness credentials of interventions that have been economically assessed using CEA frameworks. However, if studies use the same outcome measures, then their cost-effectiveness ratios can be compared.

Regarding CUA studies, there are important differences in using general health outcome measures such as DALY and QALY, whereby their results need to be interpreted with caution. This means that preventive interventions for dental caries and periodontitis cannot be directly compared or ranked against each other, nor can they be compared with other health interventions for appraisal and make health policy investment decisions. To date, there is still ongoing research to validate the use of quality-of-life survey instruments for children with dental caries, although the CHU-9D and the EQ-5D-Y instruments show promise, but these have not been tested for face validity [119]. The application of quality-of-life survey instruments for adults have had more extensive research generally [120], but they have not been widely applied in economic evaluations for preventive interventions for dental caries and periodontitis. To date, research using QALY has shown they is not sensitive to oral diseases [125], but since DALY is independent of qualityof-life survey instruments. It may be the only outcome now that can directly allow comparisons between other health interventions.

4.2 Cost-Effectiveness Analysis Methods

There were concerns regarding how the benefits were defined for CEA studies. For example, three studies (modelbased and trial-based studies) performed an economic evaluation for the cost-effectiveness of fissure sealants and fluoride varnish [40, 41], or fissure sealant and a school-based toothbrushing program [42]. In this context, fissure sealants have a clinical benefit for the occlusal surfaces of posterior molar teeth, where they are applied, whereas the clinical benefit of fluoride varnish and regular toothbrushing with fluoride toothpaste can have a spill-over benefit to other surfaces and numbers of teeth for an individual. i.e., the clinical effectiveness of each strategy was not fully incorporated and consistent.

Other studies used effectiveness data based on older studies (>20 years ago), perhaps as no recent studies were applicable [43, 44]. Other model-based studies have made assumptions for the consequences of an intervention using panel datasets of dental services, such as for the effectiveness of fissure sealants [45–47], which may or may not have a preventive effect as smooth surfaces of teeth are not protected by the fissure sealant. Similarly, three studies (modelbased and trial-based studies) investigated the cost-effectiveness of fluoride varnish interventions against standard care, where the effectiveness was evaluated for surfaces in between teeth (proximal) and not smooth surfaces [48–50]. These methodological issues on effectiveness indicate potential bias and limitation of the results.

4.3 Strengths and Limitations

One of the key strengths of the present systematic review is the breadth of the study, which includes populations of all age groups and periodontitis as one of the target oral diseases. We articulated an intervention framework to classify prevention interventions for dental caries and periodontitis according to the existing classifications for the oral disease severity. i.e., the Intervention Classification Framework for Dental Caries and Periodontitis. The framework can be useful for future systematic reviews of economic evaluations in oral health and dentistry. Given most model-based and trialbased studies were of moderate or good quality, the decision to not exclude lower quality studies is unlikely to influence the results. Method 1 from the Gonzalez-Perez (2002) study quality criteria [38] was used in preference to Methods 2 and 3 given the Drummond 10-point checklist does not include the 11th criteria question 'Is the paper relevant for the NHS and are the results sufficiently transparent for them to be replicated in an NHS setting?'

Our review expands on previous reviews by clearly articulating whether the prevention interventions were cost-effective and discussed in detail the interpretation and limitation of dental and generic health outcomes. Although dental caries is highly prevalent, their burden of disease measured by YLDs is considerably less than for periodontitis and signifies the importance for prevention efforts needed beyond dental caries.

We are aware that some publications excluded at the fulltext screening stage were of partial economic evaluations, such as the 20% sugar-sweetened beverages tax intervention [121, 122], which could be informative in the present analysis. Our review may be biased since it was restricted to publications after 2000 (inclusive), papers had to be reported in English, a limited search of databases for relevant grey literature was conducted, and relevant full-text may be excluded given the lower inter-rater reliability value of 'fair to good' agreement according to the Cohen's Kappa assessment. It is apparent that many of the studies included in our review referred to dental caries using other descriptions such as cavities and caries lesions, which could potentially result in relevant publications being excluded from our systematic review search strategy. These concerns were not observed for economic evaluations of preventive interventions for periodontitis. Finally, it is important to consider how the economic evaluation studies were validated. We did not explicitly review this as part of the a priori review protocol, and, therefore, this is essential to capture and discuss in future reviews in the field according to the ISPOR-SMDM Modeling Good Research Practices Taskforce [126].

4.4 Recommendations

We recommend three key essential considerations for future research, which can accelerate oral health policy investment: (1) use generic health outcomes to facilitate the assessment for cost-effectiveness, (2) conduct economic evaluation research to prevent periodontitis, and (3) make considerations for health equity.

5 Conclusion

Universal interventions, in particular water fluoridation, were generally demonstrated to be cost-effective, while the cost-effectiveness for selective and indicated interventions were mixed. Other universal interventions that had at least more than one study demonstrating cost-effectiveness include taxation on sugar-sweetened beverages, milk and salt fluoridation. The disparate selection of the cost components, the consequences, and the choice for measuring health outcomes, meant that it was difficult to make investment decisions for preventive interventions for dental caries and periodontitis. We propose future research on economic evaluation for the oral health context using generic health outcome measures to determine cost-effectiveness outcomes. In addition, there is an increasing need to conduct research on the cost-effectiveness of preventive interventions related to periodontitis and to reflect on health equity considerations. These recommendations will enable comparability with mainstream economic evaluations of health interventions to inform policy decision making.

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Declarations

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