

Effect of artificial community water fluoridation on dental health

Appendices



8 May 2025

Research. Evidence. Action.

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6 Appendices Question 1

6.1 Appendix A Overview of literature search results for Question 1

An initial systematic search was carried out in Medline, LILACS and Embase on the 13 July 2021. Databases such as Cochrane Library and Epistemonikos, and registers such as PROSPERO systematic review register and the Cochrane Trial Registry, were also searched for relevant material. The initial Medline and Embase searches only covered the years 1990-2021. In December 2021, on the advice of an external expert, the review team extended the date range of the review to include historical relevant material. An additional search was carried out to cover 1946–1990 using the same search strategies for the Medline and Embase databases. All material from both searches was deduplicated into Endnote, then imported into Eppi Reviewer where it was further deduplicated. This body of evidence was screened on title and abstract and then in full text. An updated search of the two main databases, using the same search strategies, was conducted on 24 February 2022, and on 28 March 2023, to capture recent evidence. The summary table below gives an overview of this complex search process (

Table 1). Search strategies for the database searches follow below.

Table 1 Overview of literature search for Q1

| Database | Date of search | Date range | No. of results |
|---|----------------|----------------------------|----------------|
| Ovid MEDLINE(R) and Epub Ahead of Print, In-Process, In-Data-Review & Other Non-Indexed Citations, Daily and Versions | 13 July 2021 | 1990- 09 July 2021 | 1,535 |
| Embase (Embase.com) | 13 July 2021 | 1990- 09 July 2021 | 2,096 |
| Cochrane Library (John Wiley & Sons Inc) | 03 Aug 2021 | November, 1946-03 Aug 2021 | 33 |
| Cochrane Trial Register (John Wiley & Sons Inc) | 03 Aug 2021 | November, 1946-03 Aug 2021 | 218 |
| Latin American and Caribbean Health Sciences Literature (LILACS) | 23 Aug 2021 | 1998-23 Aug 2021 | 23 |
| PROSPERO Trial Register | 02 Nov 2021 | Inception-02 Nov 2021 | 4 |
| Epistemonikos database of systematic reviews | 13 July 2021 | Inception-13 July 2021 | 24 |
| Ovid MEDLINE(R) and Epub Ahead of Print, In-Process, In-Data-Review & Other Non-Indexed Citations, Daily and Versions | 06 Dec 2021 | 1946- 1990 | 440 |
| Embase | 06 Dec 2021 | 1974- 1990 | 480 |
| Ovid MEDLINE(R) and Epub Ahead of Print, In-Process, In-Data-Review & Other Non-Indexed Citations, Daily and Versions | 24 Feb 2022 | 01 Dec 2021-24 Feb 2022 | 142 |
| Embase | 24 Feb 2022 | 01 Dec 2021-24 Feb 2022 | 119 |
| Ovid MEDLINE(R) and Epub Ahead of Print, In-Process, In-Data-Review & Other Non-Indexed Citations, Daily and Versions | 28 March 2032 | 01 Dec 2021-28 March 2023 | 136 |
| Embase | 28 March 2023 | 01 Dec 2021-28 March 2023 | 139 |
| Total before deduplication | | | 5,389 |
| Total after deduplication | | | 3,846 |
| Total retained for analysis after screening | | | 73 |
| Total added from reference chasing | | | 24 |
| Total included on full study and following extraction | | | 97 |

6.1.1 Medline (1990-2021)

Database(s): Ovid MEDLINE(R) and Epub Ahead of Print, In-Process, In-Data-Review & Other Non-Indexed Citations, Daily and Versions(R) 1990 to July 09, 2021

Search Strategy:

| # | Searches | Results |
|----|---|----------|
| 1 | exp Fluoridation/ | 5876 |
| 2 | Fluorides/ or Fluorine/ | 34009 |
| 3 | (fluorid* or fluorin* or flourid* or flourin* or flurid* or flurin or florid* or florin*).tw,kf,sh. and water.tw,kf,sh. /freq=5 | 1951 |
| 4 | (Hexafluorsilicic acid or Hydrofluosilicic acid or HFSA or "H2SiF6" or "CaF2" or fluorospar or fluorosilicic acid or sodium fluorosilicate\$ or silicofluorid\$).mp. | 1196 |
| 5 | or/1-4 | 39800 |
| 6 | Water Supply/ or Water/ or (drinking water or drinking suppl\$ or potable water or water suppl\$ or suppl\$ of water or public water or community water or water treatment or waterworks or water fluorid\$).mp. | 251438 |
| 7 | 5 and 6 | 6005 |
| 8 | Oral Health/ | 17919 |
| 9 | oral health.mp. and quality of life.ab,ti,hw,kw. | 5166 |
| 10 | Quality of Life/ | 216188 |
| 11 | Dental health/ | 0 |
| 12 | "Quality of Life"/ and dental health.mp. | 482 |
| 13 | (Life Quality or Health-Related Quality Of Life or Health Related Quality Of Life).mp. | 57202 |
| 14 | (Life Quality or Health-Related Quality Of Life or Health Related Quality Of Life).mp. or HRQOL.ab,ti,kw. | 57836 |
| 15 | exp Periodontal Diseases/ | 90182 |
| 16 | Dental Caries.mp. or exp Dental Caries/ | 53852 |
| 17 | carie\$.mp. | 62346 |
| 18 | Dental enamel/ | 20060 |
| 19 | exp Tooth demineralization/ | 49211 |
| 20 | ((teeth or tooth or dental or enamel or dentin or root\$) adj5 (cavit\$ or caries\$ or carious or decay\$ or lesion\$ or deminerali\$ or reminerali\$)).mp. | 86447 |
| 21 | ((teeth or tooth or dental or enamel or dentin) and plaque).mp. | 27298 |
| 22 | Tooth loss/ or tooth loss.ti,ab,hw,kw. | 7070 |
| 23 | DMF Index/ or ("DMF Index" or "Dental Plaque Index").mp. | 14647 |
| 24 | Fluorosis, dental/ or (fluorosis or fluorosed or tooth discolouration).mp. | 4311 |
| 25 | or/8-24 | 440423 |
| 26 | 7 and 25 | 2876 |
| 27 | blind*.mp. or Case-Control Studies/ or case-control.mp. or ("case series" or "time series" or "before and after").mp. or Cohort Studies/ or cohort analysis/ or cohort*.mp. or control*.mp. or clinical trial/ or Cohort Studies/ or Cross-Over Studies/ or cross over.mp. or crossover.mp. or cross sectional.mp. or double-blind method/ or doubl*.mp. or exp clinical trial/ or comparative study/ or comparative stud*.mp. or controlled clinical trial/ or Correlation study/ or cross sectional study/ or Ecological study/ or ecological stud*.mp. or ecological study.mp. or evaluation study/ or evaluation stud*.mp. or Follow-Up Studies/ or followup.mp. or follow-up.mp. or (health* adj2 survey*).mp. or Incidence/ or incidence.mp. or mask*.mp. or Mortality/ or mortality.tw. or observational study/ or Placebos/ or placebo*.mp. or predict*.mp. or Prevalence/ or prevalence.mp. or prognos*.mp. or random*.mp. or Randomized Controlled Trial/ or random allocation/ or risk.mp. or exp Research Design/ or Single-Blind Method/ or singl*.mp. or trebl*.mp. or tripl*.mp. or volunteer*.mp. | 13954657 |
| 28 | 26 and 27 | 2260 |
| 29 | limit 28 to yr="1990-Current" | 1535 |

6.1.2 Embase (1990-2021)

Database(s): Embase 1990 to 2021 July 09

Search Strategy

| # | Searches | Results |
|----|---|----------|
| 1 | Fluorides/ or Fluorine/ | 38059 |
| 2 | exp Fluoridation/ | 4376 |
| 3 | water.ti,ab,kw,sh. /freq=5 and (fluorid\$ or fluorin\$ or flourid\$ or flourin\$ or flurid\$ or flurin\$ or florid\$ or florin\$).ti,ab,kw,sh. | 2793 |
| 4 | (Hexafluorsilicic acid or Hydrofluosilicic acid or HFSA or "H2SiF6" or "CaF2" or fluorospar or fluorosilicic acid or sodium fluorosilicate\$ or silicofluorid\$).mp. | 1233 |
| 5 | or/1-4 FLUORIDE | 42694 |
| 6 | Water Supply/ or Water/ or (drinking water or drinking suppl\$ or potable water or water suppl\$ or suppl\$ of water or public water or community water or water treatment or waterworks or water fluorid\$).mp. | 426890 |
| 7 | 5 and 6 FLUORIDE AND WATER | 7651 |
| 8 | Oral Health/ | 161736 |
| 9 | oral health.mp. and quality of life.ab,ti,hw,kw. | 5379 |
| 10 | Dental health/ | 4218 |
| 11 | "Quality of Life"/ and dental health.mp. | 646 |
| 12 | Quality of Life/ | 513727 |
| 13 | (Life Quality or Health-Related Quality Of Life or Health Related Quality Of Life).mp. | 88186 |
| 14 | (Life Quality or Health-Related Quality Of Life or Health Related Quality Of Life).mp. or HRQOL.ab,ti,kw. | 90417 |
| 15 | exp Periodontal Diseases/ or periodontal disease\$.mp. | 108216 |
| 16 | exp Dental Caries/ | 49907 |
| 17 | (carie\$ or carie*).mp. | 60481 |
| 18 | ((teeth or tooth or dental or enamel or dentin or root\$) adj5 (cavit\$ or caries\$ or carious or decay\$ or lesion\$ or deminerali\$ or reminerali\$)).mp. | 71786 |
| 19 | ((teeth or tooth or dental or enamel or dentin) and plaque).mp. | 27798 |
| 20 | exp Tooth demineralization/ | 224166 |
| 21 | Fluorosis, dental/ or (fluorosis or fluorosed or tooth discolouration).mp. | 5437 |
| 22 | Dental enamel/ | 19708 |
| 23 | DMF Index/ | 961 |
| 24 | ("DMF Index" or "Dental Plaque Index").mp. | 1347 |
| 25 | Tooth loss/ or tooth loss.ti,ab,hw,kw. | 34448 |
| 26 | or/8-25 | 927462 |
| 27 | 7 and 26 ORAL HEALTH | 3443 |
| 28 | blind*.mp. or Case-Control Studies/ or case-control.mp. or ("case series" or "time series" or "before and after").mp. or Cohort Studies/ or cohort analysis/ or cohort*.mp. or control*.mp. or clinical trial/ or Cohort Studies/ or Cross-Over Studies/ or cross over.mp. or crossover.mp. or cross sectional.mp. or double-blind method/ or doubl*.mp. or exp clinical trial/ or comparative study/ or comparative stud*.mp. or controlled clinical trial/ or Correlation study/ or cross sectional study/ or Ecological study/ or ecological stud*.mp. or ecological study.mp. or evaluation study/ or evaluation stud*.mp. or Follow-Up Studies/ or followup.mp. or follow-up.mp. or (health* adj2 survey*).mp. or Incidence/ or incidence.mp. or mask*.mp. or Mortality/ or mortality.tw. or observational study/ or Placebos/ or placebo*.mp. or predict*.mp. or Prevalence/ or prevalence.mp. or prognos*.mp. or random*.mp. or Randomized Controlled Trial/ or random allocation/ or risk.mp. or exp Research Design/ or Single-Blind Method/ or singl*.mp. or trebl*.mp. or tripl*.mp. or volunteer*.mp. | 21777714 |
| 29 | 27 and 28 FLUROIDE + WATER + ORAL HEALTH | 2508 |
| 30 | limit 29 to yr="1990-Current" | 2096 |

6.1.3 Cochrane Central (1946-03 August 2021)

| # | Searches | Results |
|--------------------|--|---------|
| #1 | MeSH descriptor: [Fluorides] explode all trees | 2686 |
| #2 | MeSH descriptor: [Fluorine] explode all trees | 85 |
| #3 | MeSH descriptor: [Fluoridation] explode all trees | 38 |
| #4 | ((fluorid* or fluorin* or flurin* or flurid* or flourid* or flourin*)) | 6295 |
| #5 | #1 or #2 or #3 or #4 | 6375 |
| #6 | MeSH descriptor: [Water Supply] explode all trees | 180 |
| #7 | MeSH descriptor: [Water] explode all trees | 2429 |
| #8 | ("water treatment") | 320 |
| #9 | water near fluorid* | 255 |
| #10 | ("community water" OR "community-based water" OR "community supply" OR "community fluoridation") | 28 |
| #11 | #6 OR #7 OR #8 OR #9 OR #10 | 3044 |
| #12 | #5 and #11 | 306 |
| #13 | MeSH descriptor: [Oral Health] explode all trees | 451 |
| #14 | MeSH descriptor: [Tooth Diseases] explode all trees | 11401 |
| #15 | MeSH descriptor: [DMF Index] explode all trees | 518 |
| #16 | MeSH descriptor: [Dental Enamel] explode all trees | 1168 |
| #17 | #13 OR #14 OR #15 OR #16 | 12063 |
| #18 | ("oral health" OR "dental health"):ti,ab,kw | 3921 |
| #19 | (caries OR carious OR cavit* OR decay* OR demineral* OR remineral* OR "dental plaque index") | 22874 |
| #20 | (fluorosis or fluorosed OR ((tooth OR teeth) NEXT (discolour* OR discolor*))) | 784 |
| #21 | (enamel OR root OR dentin OR tooth OR teeth OR oral OR dental):ti,ab,kw | 209510 |
| #22 | ("quality of life" OR "life quality" OR QoL OR HRQoL):ti,ab,kw | 124121 |
| #23 | #21 and #22 | 13571 |
| #24 | #17 OR #18 OR #19 OR #20 OR #23 | 43597 |
| #25 | #12 and #24 | 251 |
| Cochrane Reviews | | 22 |
| Cochrane Protocols | | 2 |

| # | Searches | Results |
|------------------|----------|---------|
| Trials | | 218 |
| Clinical Answers | | 9 |

6.1.4 Latin American and Caribbean Health Sciences Literature Database (LILACS) (1998-23 August 2021)

| # | Searches | Result |
|--------|--|--------|
| #1 | (fluoride OR fluorine OR fluori\$ OR fluoruro OR fluoreto) AND | |
| #2 | ((teeth OR tooth OR dental OR dentin\$ OR enamel OR root\$) OR (Cavit\$ OR caries OR carious OR decay*)) AND | |
| Total: | | 23 |

6.1.5 Epistemonikos (2009-13 August 2021)

| # | Searches | Result |
|--------|---|--------|
| | (title:((title:(fluoride AND water fluoridation AND review) OR abstract:(fluoride AND water fluoridation AND review)) AND (teeth OR tooth OR dental OR dentin* OR enamel OR root*) OR (cavit* OR caries OR carious OR decay*)) OR abstract:((title:(fluoride AND water fluoridation AND review) OR abstract:(fluoride AND water fluoridation AND review)) AND (teeth OR tooth OR dental OR dentin* OR enamel OR root*) OR (cavit* OR caries OR carious OR decay*))) | |
| Total: | | 24 |

6.1.6 Supplementary grey literature table

General scoping searches for Question 1 were carried out in the search engine *Google.com* to gain an initial idea of terminology and likely key terms. Reviewing literature and systematic reviews in the area (retrieved in the Epistemonikos and Cochrane databases) also helped build up our search vocabulary. Search terms used included combinations of water, fluoridated water, fluoride, oral health, dental health; For Questions 2A and 2B, vocabulary around children and topical fluoride(s) was added. Broad terms were used, in structured searches, to capture as much relevant material as possible. Further searches were carried out using the websites of relevant bodies. Results were pre-screened by the information specialist (AF) and screened using different screening codes for questions 1, 2A and 2B (OC). Updated searches of these resources was undertaken in March, 2023, as well as the search engine, *DuckDuckGo*. A full list of grey literature resources is available at Table 2.

Table 2 Structured grey literature search

| Country | Body | Link | Date | Search string | Limit | # |
|------------------|--|---|-----------|---|----------------|----|
| Australia | | | | | | |
| | Australian Dental Association (ADA) | https://www.ada.org.au/Dental-Professionals | 09-Feb-22 | fluoride and water and "oral health" | Publications | 21 |
| | | | | fluoride and water and "oral health" and children | Publications | 22 |
| | Fluoride Reference Group | https://www.nhmrc.gov.au/about-us/leadership-and-governance/committees/fluoride-reference-group | 09-Feb-22 | | | 3 |
| | National Health and Medical Research Council (NHMRC) | https://www.nhmrc.gov.au/ | 09-Feb-22 | fluoride and water and "oral health" | | 10 |
| | | | | fluoride and children | | 12 |
| Canada | | | | | | |
| | Canadian Dental Association | https://www.cda-adc.ca/en/index.asp | 09-Feb-22 | fluoride and water and "oral health" and children | | 3 |
| | Canadian Institute for Health Information | https://www.cihi.ca/en | 09-Feb-22 | fluoride and water and "oral health" | | 0 |
| | | | | fluoride and water and "oral health" and children | | 0 |
| | Health Canada | https://www.canada.ca/en/health-canada.html | 09-Feb-22 | Fluorosis | | 0 |
| | | | | Flu* | | 0 |
| | | | | Fluoride and water and | Terms in title | 26 |

| Country | Body | Link | Date | Search string | Limit | # |
|--------------------|---|---|-----------|---|---|----|
| | | | | "oral health" | | |
| Ireland | | | | | | |
| | Dental Council (Ireland) | http://www.dentalcouncil.ie | 09-Feb-22 | No search function on website. Nothing relevant retrieved via browsing. | | 0 |
| | Department of Health | https://www.health.gov.ie | 28-Feb-22 | fluoride and water and "oral health" | Policies/Policy Information / Publications / Reports/ | 15 |
| | Irish Expert Body on Fluorides and Health | https://www.fluoridesandhealth.ie/ | 28-Feb-22 | fluoride and | | 9 |
| | | | | water fluoridation and "oral health" and "topical fluorides" and children | | 45 |
| | | | | water fluoridation and "oral health" and topical | | 19 |
| | Irish Dental Association | https://www.dentist.ie/ | 28-Feb-22 | water fluoridation and "oral health" and topical | | 19 |
| | | | | fluoride and water and "oral health" and children | | 0 |
| | | | | fluoride and water | | 0 |
| | | | | Fluoride | | 11 |
| New Zealand | | | | | | |
| | Environmental Health Intelligence New Zealand (EHINZ) | https://www.ehinz.ac.nz/ | 28-Feb-22 | water + fluoride + children | | 1 |

| Country | Body | Link | Date | Search string | Limit | # |
|-----------|---------------------------------------|---|-------------|---|--|----|
| | Ministry of Health | https://www.health.govt.nz/ | | | | |
| | | | 28-Feb-22 | fluoride and water and "oral health" | | 54 |
| | | | | fluoride and water and "oral health" and topical and children | Publication type: strategies and plans; evaluation and review; statistical publications ; guides and standards | 5 |
| | New Zealand Dental Association (NZDA) | https://www.nzda.org.nz/ | 09-Feb-22 | fluoride and water and "oral health" | | 31 |
| UK | | | | | | |
| | British Dental Association (BDA) | https://www.bda.org/ | 09-Feb-22 | fluoride and water and "oral health" | | 31 |
| | Department of Health | https://www.gov.uk/government/organisations/department-of-health-and-social-care | 22 March 22 | fluoride and water and "oral health" | "Guidance and regulation" AND Limiter: "Research and statistics" AND "policy papers and consultations" AND Topic: "Health and Social Care" | 82 |
| | | | | fluoride and water and "oral health" and children and topical | "Guidance and regulation" AND Limiter: "Research and statistics" AND "policy papers and | 70 |

| Country | Body | Link | Date | Search string | Limit | # |
|---------|---|---|-------------|---|--|----|
| | | | | | consultations" AND Topic: "Health and Social Care" | |
| | National Institute for Health and Care Excellence (NICE) | https://www.nice.org.uk/ | 28-Feb-22 | water+fluoride | | 4 |
| | | | | children+fluoride | | 13 |
| | Scottish Dental Clinical Effectiveness Programme | https://www.sdcep.org.uk/?s=fluoridation+or+fluoride | 28-Feb-22 | children+fluoride | | 17 |
| USA | | | | | | |
| | American Dental Association (ADA) | https://www.ada.org/ | 09-Feb-22 | fluoride and water and "oral health" | article+ JADA+ Clinical Research+ Fluoride+ Regulatory+ Science & Technology | 42 |
| | American Association of Pediatric Dentistry (AAPD) | https://www.aapd.org/ | 28-Feb-22 | Fluoride and water and "oral health" and topical | Educational resources (no access to any other material) | 27 |
| | American Academy of Oral Medicine | https://www.aaom.com | 28-Feb-22 | Fluoride | | 31 |
| | Centers for Disease Control (CDC) | https://www.cdc.gov/fluoridation/ | 28-Feb-2022 | "Water fluoridation" and "oral health" and "topical fluoride" | | 53 |
| | Centers for Disease Control (CDC) Community Water Fluorosis | https://www.cdc.gov/fluoridation/faqs/dental_fluorosis/ | 28-Feb-2022 | | | 1 |
| | Centers for Disease Control (CDC) | https://www.cdc.gov/fluoridation/basics/fluoride- | 28-Feb-2022 | | | 1 |

| Country | Body | Link | Date | Search string | Limit | # |
|----------------------|--|---|-----------|---|----------|-----|
| | Fluoride Products | products.html | | | | |
| | Dept of Health and Human Services (HHS) | https://www.hhs.gov/ | 28-Feb-22 | fluoride and water and "oral health" and children and topical | | 0 |
| International | | | | | | |
| | Canada's Drug & Health Technology Agency (CADTH) | https://www.cadth.ca/grey-matters-practical-tool-searching-health-related-grey-literature-0 | 28-Feb-22 | | | 10 |
| | Centre for Evidence-based Dentistry | https://www.cebd.org/ | 28-Feb-22 | Fluoride and water | | |
| | Core.Ac.uk | https://core.ac.uk/ | 28-Feb-22 | children and fluoride | | |
| | Council of European Dentists (CED) | https://www.eudental.eu/ | 28-Feb-22 | Fluoride and oral health | research | 124 |
| | European Food Safety Authority | https://www.efsa.europa.eu/en | 28-Feb-22 | fluoridated water and "oral health" and "topical fluoride" and children | research | 31 |
| | Health Systems Evidence | https://www.healthsystemsevidence.org/ | 28-Feb-22 | water and fluoride and oral health | | 0 |
| | International Association for Dental Research (IADR) | https://www.iadr.org/ | 28-Feb-22 | | | |
| | International Network of Agencies for Health | https://database.inahta.org/about#about-inahta | 28-Feb-22 | water fluoridation and dental and children | | 2 |

| Country | Body | Link | Date | Search string | Limit | # |
|---------|--------------------------------|---|-----------|---|------------------------------|-------|
| | Technology Assessment (INAHTA) | | | | | |
| | Med Archives | https://www.medrxiv.org/ | 28-Feb-22 | "water and fluoride and dental" | | 14 |
| | World Dental Federation (FDI) | https://www.fdiworlddental.org/ | 28-Feb-22 | carious OR caries Or Cari* + fluoridation (condition) AND fluoridation OR fluoride (intervention) | All Phases + Children trials | 0 |
| Total: | | | | | | 1,013 |

6.1.7 Overview of updated database search (23 Feb 2023)

An updated database search was performed in the main databases (Medline, Embase) to capture recent relevant data using the same search strategy as the original search to the current date. See

Table 1.

6.2 Appendix B PRISMA checklist and PRISMA-S for Question 1

6.2.1 PRISMA checklist for Question 1

| Topic | Item | Checklist item | Location where item is reported |
|-------------------------|------|--|---|
| TITLE | | | |
| Title | 1 | Identify the report as a systematic review. | Title page |
| ABSTRACT | | | |
| Abstract | 2 | See the PRISMA 2020 for Abstracts checklist. | Executive summary |
| INTRODUCTION | | | |
| Rationale | 3 | Describe the rationale for the review in the context of existing knowledge. | Section 1.1.4 |
| Objectives | 4 | Provide an explicit statement of the objective(s) or question(s) the review addresses. | Section 1.2 |
| METHODS | | | |
| Eligibility criteria | 5 | Specify the inclusion and exclusion criteria for the review and how studies were grouped for the syntheses. | Sections 2.3.1 |
| Information sources | 6 | Specify all databases, registers, websites, organisations, reference lists and other sources searched or consulted to identify studies. Specify the date when each source was last searched or consulted. | Sections 2.4 |
| Search strategy | 7 | Present the full search strategies for all databases, registers and websites, including any filters and limits used. | Appendix A of Section 6 |
| Selection process | 8 | Specify the methods used to decide whether a study met the inclusion criteria of the review, including how many reviewers screened each record and each report retrieved, whether they worked independently, and if applicable, details of automation tools used in the process. | Section 2.6 Error! Reference source not found. |
| Data collection process | 9 | Specify the methods used to collect data from reports, including how many reviewers collected data from each report, whether they worked independently, any processes for obtaining or confirming data from study investigators, and if applicable, details of automation tools used in the process. | Section 2.7 |

| Topic | Item | Checklist item | Location where item is reported |
|-------------------------------|------|---|--|
| Data items | 10a | List and define all outcomes for which data were sought. Specify whether all results that were compatible with each outcome domain in each study were sought (e.g. for all measures, time points, analyses), and if not, the methods used to decide which results to collect. | Sections 2.3.1, Table 2, Section 2.7.1.1 and 2.7.1.2 |
| | 10b | List and define all other variables for which data were sought (e.g. participant and intervention characteristics, funding sources). Describe any assumptions made about any missing or unclear information. | Section 2.7, 2.7.1.1 and 2.7.1.2 |
| Study risk of bias assessment | 11 | Specify the methods used to assess risk of bias in the included studies, including details of the tool(s) used, how many reviewers assessed each study and whether they worked independently, and if applicable, details of automation tools used in the process. | Section 2.8 |
| Effect measures | 12 | Specify for each outcome the effect measure(s) (e.g. risk ratio, mean difference) used in the synthesis or presentation of results. | Section 2.9.1 |
| Synthesis methods | 13a | Describe the processes used to decide which studies were eligible for each synthesis (e.g. tabulating the study intervention characteristics and comparing against the planned groups for each synthesis (item #5)). | Section 2.9.2 |
| | 13b | Describe any methods required to prepare the data for presentation or synthesis, such as handling of missing summary statistics, or data conversions. | Section 2.9.3 |
| | 13c | Describe any methods used to tabulate or visually display results of individual studies and syntheses. | Section 2.9.2 and 2.9.3 |
| | 13d | Describe any methods used to synthesize results and provide a rationale for the choice(s). If meta-analysis was performed, describe the model(s), method(s) to identify the presence and extent of statistical heterogeneity, and software package(s) used. | Section 2.9.3 |
| | 13e | Describe any methods used to explore possible causes of heterogeneity among study results (e.g. subgroup analysis, meta-regression). | Section 2.9.3 |
| | 13f | Describe any sensitivity analyses conducted to assess robustness of the synthesized results. | Section 2.9.3 |
| Reporting bias assessment | 14 | Describe any methods used to assess risk of bias due to missing results in a synthesis (arising from reporting biases). | Section 2.9.2 and 2.9.3 |

| Topic | Item | Checklist item | Location where item is reported |
|-------------------------------|------|--|--|
| Certainty assessment | 15 | Describe any methods used to assess certainty (or confidence) in the body of evidence for an outcome. | Section 2.10 |
| RESULTS | | | |
| Study selection | 16a | Describe the results of the search and selection process, from the number of records identified in the search to the number of studies included in the review, ideally using a flow diagram. | Section 3.1.1, Appendix F of Section 6 |
| | 16b | Cite studies that might appear to meet the inclusion criteria, but which were excluded, and explain why they were excluded. | Section 3.1.4.1, Tables 11 and 12, Section 3.1.4.2, Tables 13 and 14 Appendix C of Section 6 |
| Study characteristics | 17 | Cite each included study and present its characteristics. | Section 3.1.2, Table 8, Section 3.1.5, Table 36 |
| Risk of bias in studies | 18 | Present assessments of risk of bias [quality assessment] for each included study. | Appendix H of Section 6 |
| Results of individual studies | 19 | For all outcomes, present, for each study: (a) summary statistics for each group (where appropriate) and (b) an effect estimate and its precision (e.g. confidence/credible interval), ideally using structured tables or plots. | Section 3.1.4.3, Tables 15–35, Section 3.1.7, Section 3.1.7.4, Tables 38 – 44, Appendix J of Section 6 |
| Results of syntheses | 20a | For each synthesis, briefly summarise the characteristics and risk of bias among contributing studies. | Sections 3.1.3, Table 9 and 10, Section 3.1.6, Table 37, Appendix H of Section 6 |
| | 20b | Present results of all statistical syntheses conducted. If meta-analysis was done, present for each the summary estimate and its precision (e.g. confidence/credible interval) and measures of statistical heterogeneity. If comparing groups, describe the direction of the effect. | Section 3.1.4.4, Figures 4 -8, Section 3.1.7 |

| Topic | Item | Checklist item | Location where item is reported |
|---------------------------|------|--|---|
| Reporting biases | 20c | Present results of all investigations of possible causes of heterogeneity among study results. | 3.1.4.4 |
| | 20d | Present results of all sensitivity analyses conducted to assess the robustness of the synthesized results. | Appendix K of Section 6 |
| | 21 | Present assessments of risk of bias due to missing results (arising from reporting biases) for each synthesis assessed. | Not applicable as mainly cross section surveys |
| Certainty of evidence | 22 | Present assessments of certainty (or confidence) in the body of evidence for each outcome assessed. | Appendix L of Section 6 |
| DISCUSSION | | | |
| Discussion | 23a | Provide a general interpretation of the results in the context of other evidence. | Sections 4.1 and 4.2 |
| | 23b | Discuss any limitations of the evidence included in the review. | Section 4.3 |
| | 23c | Discuss any limitations of the review processes used. | Section 4.3 |
| | 23d | Discuss implications of the results for practice, policy, and future research. | Sections 4.4 and 4.5 |
| OTHER INFORMATION | | | |
| Registration and protocol | 24a | Provide registration information for the review, including register name and registration number, or state that the review was not registered. | Section 2.3.2 |
| | 24b | Indicate where the review protocol can be accessed, or state that a protocol was not prepared. | Section 2.3.2 |
| | 24c | Describe and explain any amendments to information provided at registration or in the protocol. | Section 2.3.2 |
| Support | 25 | Describe sources of financial or non-financial support for the review, and the role of the funders or sponsors in the review. | Not applicable as all authors are salaried public servants who are funded from the DOH public funding and are obliged to be objective |
| Competing interests | 26 | Declare any competing interests of review authors. | None |

| Topic | Item | Checklist item | Location where item is reported |
|--|------|--|----------------------------------|
| Availability of data, code and other materials | 27 | Report which of the following are publicly available and where they can be found: template data collection forms; data extracted from included studies; data used for all analyses; analytic code; any other materials used in the review. | Appendix D, E and J of Section 6 |

Source: Page *et al.* (2021)[28]

6.2.2 PRISMA-S for Question 1

| Section/topic | # | Checklist item | Location(s) Reported |
|--|----|--|--|
| INFORMATION SOURCES AND METHODS | | | |
| Database name | 1 | Name each individual database searched, stating the platform for each. | Sections 2.4.4 and 2.5 |
| Multi-database searching | 2 | If databases were searched simultaneously on a single platform, state the name of the platform, listing all of the databases searched. | n/a |
| Study registries | 3 | List any study registries searched. | Sections 2.4 and 2.5 |
| Online resources and browsing | 4 | Describe any online or print source purposefully searched or browsed (e.g., tables of contents, print conference proceedings, web sites), and how this was done. | Sections 2.4.5 and 2.4.6 |
| Citation searching | 5 | Indicate whether cited references or citing references were examined, and describe any methods used for locating cited/citing references (e.g., browsing reference lists, using a citation index, setting up email alerts for references citing included studies). | Section 2.4.5 |
| Contacts | 6 | Indicate whether additional studies or data were sought by contacting authors, experts, manufacturers, or others. | n/a |
| Other methods | 7 | Describe any additional information sources or search methods used. | n/a |
| SEARCH STRATEGIES | | | |
| Full search strategies | 8 | Include the search strategies for each database and information source, copied and pasted exactly as run. | Appendix A of Section 6 |
| Limits and restrictions | 9 | Specify that no limits were used, or describe any limits or restrictions applied to a search (e.g., date or time period, language, study design) and provide justification for their use. | Sections 2.5 and Appendix A of Section 6 |
| Search filters | 10 | Indicate whether published search filters were used (as originally designed or modified), and if so, cite the filter(s) used. | n/a |
| Prior work | 11 | Indicate when search strategies from other literature reviews were adapted or reused for a substantive part or all of the search, citing the previous review(s). | n/a |

| | | | |
|-------------------------|--------|--|-------------------------|
| Updates | 1 2 | Report the methods used to update the search(es) (e.g., rerunning searches, email alerts). | Appendix A of Section 6 |
| Dates of searches | 1 3 | For each search strategy, provide the date when the last search occurred. | Appendix A of Section 6 |
| PEER REVIEW | | | |
| Peer review | 1 4 | Describe any search peer review process. | Section 2.4.3 |
| MANAGING RECORDS | | | |
| Total Records | 1 5 | Document the total number of records identified from each database and other information sources. | Appendix A of Section 6 |
| Deduplication | 1 6 | Describe the processes and any software used to deduplicate records from multiple database searches and other information sources. | Section 2.4.4 |

PRISMA-S: An Extension to the PRISMA Statement for Reporting Literature Searches in Systematic Reviews

Rethlefsen ML, Kirtley S, Waffenschmidt S, Ayala AP, Moher D, Page MJ, Koffel JB, PRISMA-S Group.

Last updated February 27, 2020.

6.3 Appendix C Studies excluded at full text and extraction screening stages by reason for exclusion

6.3.1 Exclude on population

Exclude on population (n=4)

Hobbs M, Wade A, Jones P, *et al.* Area-level deprivation, childhood dental ambulatory sensitive hospitalizations and community water fluoridation: evidence from New Zealand. *Int J Epidemiol* 2020;49:908–16. <https://doi.org/10.1093/ije/dyaa043>

Nunn JH, Ekanayake L, Rugg-Gunn AJ, *et al.* Assessment of enamel opacities in children in Sri Lanka and England using a photographic method. *Community Dent Health* 1993;10:175–88.

Takeuchi K, Nakagaki H, Toyama Y, *et al.* Fluoride concentrations and distribution in premolars of children from low and optimal fluoride areas. *Caries Res* 1996;30:76–82.

Comparing the effects of milk and soy-based drinks on tooth enamel. CN-01970673. *Cochrane Cent Regist Control Trials* 2019;9.

<https://www.cochranelibrary.com/central/doi/10.1002/central/CN-01970673/full>

6.3.2 Exclude on intervention

Exclude on intervention (n=177)

Abramson E. The fluoride content in drinking water and the incidence of dental caries among Swedish school-children of eight years. *Odontol Tidskr* 1955;62:493–7.

Adair S, Hanes C, Russell C, *et al.* Dental caries and fluorosis among children in a rural Georgia area. *Pediatr Dent* 1999;21:81–5.

Adler P, Straub J. A water-borne caries-protective agent other than fluorine. *Acta Med Acad Sci Hung* 1953;4:221–7.

Ahmed AT, Soto-Rojas A, Dean J, *et al.* Demarcated Primary Second Molar Hypomineralization: Prevalence Data and Associated Sociodemographic Determinants from Indiana. *Pediatr Dent* 2021;43:443–50.

Ainamo J, Parvianinen K. Influence of increased toothbrushing frequency on dental health in low, optimal, and high fluoride areas in Finland. *Community Dent Oral Epidemiol* 1989;17:296–9. doi:10.1111/j.1600-0528.1989.tb00640.x

Anderson L, Martin NR, Flynn RT, *et al.* The importance of substate surveillance in detection of geographic oral health inequalities in a small state. *J Public Health Manag Pract JPHMP* 2012;18:461–8. doi:10.1097/PHH.0b013e31825eabbb

Andrik P, Muncnerova Z. [Effect of low contents of fluorides in drinking water on the incidence of dental caries]. *Bratisl Lek Listy* 1955;35:154–62.

Antunes JLF, Narvai PC, Nugent ZJ. Measuring inequalities in the distribution of dental caries. *Community Dent Oral Epidemiol* 2004;32:41–8. doi:10.1111/j.1600-0528.2004.00125.x

Antunes JLF, Peres MA, de Campos Mello TR, *et al.* Multilevel assessment of determinants of dental caries experience in Brazil. *Community Dent Oral Epidemiol* 2006;34:146–52. doi:10.1111/j.1600-0528.2006.00274.x

Ardenghi TM, Piovesan C, Antunes JLF. Desigualdades na prevalência de carie dentária não tratada em crianças pré-escolares no Brasil. *Rev Saúde Pública* 2013;47:129–37.

Armfield JM. Public water fluoridation and dental health in New South Wales. *Aust N Z J Public Health* 2005;29:477–83. doi:10.1111/j.1467-842X.2005.tb00230.x

Exclude on intervention (n=177)

- Armfield JM. The benefits of water fluoridation across areas of differing socio-economic status. *Aust Dent J* 2008;53:180–3. doi:10.1111/j.1834-7819.2008.00030.x
- Armfield JM. Community effectiveness of public water fluoridation in reducing children's dental disease. *Public Health Rep* 2010;125:655–64. doi:10.1177/003335491012500507
- Armfield JM, Spencer AJ. Consumption of nonpublic water: implications for children's caries experience. *Community Dent Oral Epidemiol* 2004;32:283–96. doi:10.1111/j.1600-0528.2004.00167.x
- Armfield JM, Spencer AJ, Roberts-Thomson KF, *et al.* Water fluoridation and the association of sugar-sweetened beverage consumption and dental caries in Australian children. *Am J Public Health* 2013;103:494–500. doi:10.2105/AJPH.2012.300889
- Arnold FA. Grand Rapids fluoridation study; results pertaining to the eleventh year of fluoridation. *Am J Public Health Nations Health* 1957;47:539–45. doi:10.2105/ajph.47.5.539
- Arora A, Hawks RW. Dental caries in children: a comparison of one non-fluoridated and two fluoridated communities in NSW. *New South Wales Public Health Bull* 2010;21:257–62. doi:10.1071/nb10029
- Arora S, Kumar JV, Moss ME. Does water fluoridation affect the prevalence of enamel fluorosis differently among racial and ethnic groups? *J Public Health Dent* 2018;78:95–9. doi:10.1111/jphd.12258
- Arrow P, Piggott S, Carter S, *et al.* Atraumatic Restorative Treatments in Australian Aboriginal Communities: A Cluster-randomized Trial. *JDR Clin Transl Res* 2021;6:430–9. doi:10.1177/2380084420963949
- Attwood D, Blinkhorn AS. A reassessment of the dental health of urban Scottish schoolchildren following the cessation of water fluoridation. *Community Dent Health* 1989;6:207–14.
- Axelsson P, Buischi YA, Barbosa MF, *et al.* The effect of a new oral hygiene training program on approximal caries in 12-15-year-old Brazilian children: results after three years. *Adv Dent Res* 1994;8:278-284. doi:10.1177/08959374940080022201
- Azcurra AI, Battellino LJ, Calamari SE, *et al.* Estado de salud bucodental de escolares residentes en localidades abastecidas con agua de consumo humano de muy alto y muy bajo contenido de fluoruros. *Rev Saúde Pública* 1995;29:364–75.
- Baldani MH, Narvai PC, Antunes JLF. Dental caries and socioeconomic conditions in the State of Paraná, Brazil, 1996. *Cad Saude Publica* 2002;18:755–63.
- Barrett MJ, Williamson JJ. Oral health of Australian aborigines: survey methods and prevalence of dental caries. *Aust Dent J* 1972;17:37–50.
- Batsos C, Boyes R, Mahar A. Community water fluoridation exposure and dental caries experience in newly enrolled members of the Canadian Armed Forces 2006-2017. *Can J Public Health* 2021;112:513–20. doi:10.17269/s41997-020-00463-7
- Bellinger W, Mankin JD. Effect of controlled fluoridated public water supplies on the dental caries experience for children ages 9 through 12 in three Kansas cities. *J Kans State Dent Assoc* 1965;49:117–20.
- Beltran-Aguilar ED, Barker L, Dye BA. Prevalence and severity of dental fluorosis in the United States, 1999-2004. *NCHS Data Brief* 2010;;1–8.
- Binder K. Comparison of the effects of fluoride drinking water on caries frequency and mottled enamel in three similar regions of Austria over a 10-year period. *Caries Res* 1973;7:179–83.
- Blayney JR. A report on thirteen years of water fluoridation in Evanston, Ill. *J Am Dent Assoc* 1960;61:76–9.
- Bohaty BS, Parker WA, Seale NS, *et al.* The prevalence of fluorosis-like lesions associated with topical and systemic fluoride usage in an area of optimal water fluoridation. *Pediatr Dent* 1989;11:125–8.

Exclude on intervention (n=177)

Bomfim Rafael Aiello, Frazao Paulo. Impact of water fluoridation on dental caries decline across racial and income subgroups of Brazilian adolescents. *Epidemiol Health* 2022;44:e2022007. doi:10.4178/epih.e2022007

Bomfim RA, Frias AC, Cascaes AM, *et al.* Sedentary behavior, unhealthy food consumption and dental caries in 12-year-old schoolchildren: a population-based study. *Braz Oral Res* 2021;35:e041. doi:10.1590/1807-3107bor-2021.vol35.0041

Broffitt B, Levy SM, Warren J, *et al.* Factors associated with surface-level caries incidence in children aged 9 to 13: The Iowa Fluoride Study. *J Public Health Dent* 2013;73:304–10. doi:10.1111/jphd.12028

Broffitt B, Marshall TA, Eichenberger-Gilmore JM, *et al.* Associations between fluorosis of permanent incisors and fluoride intake from infant formula, other dietary sources and dentifrice during early childhood. *J Am Dent Assoc* 2010;141:1190–201. doi:10.14219/jada.archive.2010.0046

Bronson ME. Dental health in an area of maximum water fluoridation. *Dent Hyg (Chic)* 1982;56:38–41.

Brun C, Thylstrup A. Fluoride in whole saliva and dental caries experience in areas with high or low concentrations of fluoride in the drinking water. *Caries Res* 1984;18:450–6.

Butler WJ, Segreto V, Collins E. Prevalence of dental mottling in school-aged lifetime residents of 16 Texas communities. *Am J Public Health* 1985;75:1408–12.

Cabral RN, Leal SC, Bernardino Í de M, *et al.* Caries lesion transition patterns of schoolchildren in a fluoridated community in Brazil. *Clin Oral Investig* 2022;26:689–95. doi:10.1007/s00784-021-04046-9

Celeste RK, Nadanovsky P, De Leon AP. [Association between preventive care provided in public dental services and caries prevalence]. *Rev Saude Publica* 2007;41:830–8. doi:10.1590/s0034-89102007000500018

Cisternas P, Guerrero S, Morales A, *et al.* Dietary ingestion of fluoride and caries prevalence in preschool and school children in cities with different fluoride content in the drinking water and diet. *Rev Med Chil* 1994;122:459–64.

Clark DC, Berkowitz J. The relationship between the number of sound, decayed, and filled permanent tooth surfaces and the number of sealed surfaces in children and adolescents. *J Public Health Dent* 1997;57:171–5. doi:10.1111/j.1752-7325.1997.tb02969.x

Clune TW. Prevalence of dental caries in primary and permanent teeth; a study of 8,853 school children. *R I Med J* 1953;36:653–7; *passim*.

Colquhoun J. The influence of social rank and fluoridation on dental treatment requirements. *N Z Dent J* 1977;73:146–8.

Colquhoun J. Influence of social class and fluoridation on child dental health. *Community Dent Oral Epidemiol* 1985;13:37–41.

Correia Sampaio F, Ramm von der Fehr F, Arneberg P, *et al.* Dental fluorosis and nutritional status of 6- to 11-year-old children living in rural areas of Paraíba, Brazil. *Caries Res* 1999;33:66–73. doi:10.1159/000016497

Crall JJ, Edelstein B, Tinanoff N. Relationship of microbiological, social, and environmental variables to caries status in young children. *Pediatr Dent* 1990;12:233–6.

Curzon ME, Richardson DS, Featherstone JD. Dental caries prevalence in Texas schoolchildren using water supplies with high and low lithium and fluoride. *J Dent Res* 1986;65:421–3.

Cutress TW, Coote GE, Shu M, *et al.* Fluoride content of the enamel and dentine of human premolars prior to and following the introduction of fluoridation in New Zealand. *Caries Res* 1996;30:204–12.

Dalla Nora Â, Dalmolin A, Gindri LD, *et al.* Oral health status of schoolchildren living in rural and urban areas in southern Brazil. *Braz Oral Res* 2020;34:e060. doi:10.1590/1807-3107bor-2020.vol34.0060

de Liefde B, Ritchie GR. Evaluation in dental public health in New Zealand. *N Z Dent J* 1984;80:8–14.

Exclude on intervention (n=177)

Dean HT, Arnold FA, Jay P, *et al.* Studies on mass control of dental caries through fluoridation of the public water supply. *Public Health Rep Wash DC* 1896 1950;65:1403–8.

Deatherage CF. Fluoride Domestic Waters and Dental Caries Experience in 2026 White Illinois Selective Service Men. *J Dent Res* 1943;22:129–37. doi:10.1177/00220345430220020501

Del Cid C. [Natural fluorides and incidence of caries in the province of San Juan, Republic of Argentina] Fluor natural y prevalencia de caries en la provincia de San Juan--Republica Argentina. *Rev Asoc Odontol Argent* 1967;55:382–6.

Department of Health. Healthy Ireland Survey 2018. 2018. <https://www.gov.ie/en/publication/612079-healthy-ireland-survey-2018/> (accessed 19 Jun 2023).

Devoto FC, Bordoni NE, De Manfredi CF. Dental caries in deciduous teeth of nineteenth century Araucanians. *J Dent Res* 1968;47:571–4.

Do LG, Ha DH, Roberts-Thomson KF, *et al.* Race- and Income-Related Inequalities in Oral Health in Australian Children by Fluoridation Status. *JDR Clin Transl Res* 2018;3:170–9. doi:10.1177/2380084417751350

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6.3.8 Exclude unobtainable

Exclude – unobtainable (n=8)

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6.4 Appendix D Extraction forms

Table 3 General information form

| Study ID From Eppi | Author First author | Year Of publication | Location County | Area State/County/City/Town | Objective Aim of study | Secondary publication Data will not be extracted unless additional endpoints | Associated papers Same overall project differences | Study design HRB decision | Participant age Mean or ranges described in study | Artificial fluoridation Confirm if explicitly stated (Y/N) | Fluoride interventions | Outcome Oral health outcome assessed | Outcome details Including method of measurement | Extracted | Validated |
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Table 4 Study design and implementation form

| Study ID | Author | Year | Study design (Author allocated) | Study design (HRB allocated) | Justification | Length of study | Length of exposure to CWF | Details of exposure | Details of comparator | Eligibility criteria | Sample size calculation | Response rate | Blinding of assessors to exposure | % Lost to follow-up | Method for handling missing data | Data collection | Confounders | Control for confounding | Identification of effect modification | Effect modifiers | Notes |
|---------------------------|---------------------|-----------------------|---------------------------------|-----------------------------------|---------------|-----------------|---|-----------------------|-----------------------|----------------------|---|---------------|-----------------------------------|--------------------------|--|--------------------------|-------------|-------------------------|---------------------------------------|------------------|-------|
| Form Epi | <i>First author</i> | <i>Of publication</i> | <i>As stated in the study</i> | <i>As agreed by research team</i> | | | <i>Length of time exposed to community water fluoridation</i> | <i>Including dose</i> | <i>Including dose</i> | | <i>expected prevalence, power to detect a difference and allowed variance, results CIs calculated</i> | | | <i>For main analysis</i> | <i>e.g. last observation carried forward</i> | <i>Brief description</i> | | | <i>Yes or not reported</i> | | |
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Table 5 Study participants characteristics form

| Study ID | Author | Year | Group for characteristics | N | Mean age/Age range | % Female | N included in final analysis |
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| <i>From Eppi</i> | <i>First author</i> | <i>Of publication</i> | | <i>Enrolled</i> | | | |
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Table 6 Study measurement and outcomes form

| Of publication | Outcome of interest: Caries | % caries primary teeth | % caries free primary teeth | % caries permanent teeth | % caries free permanent teeth | dmft/deft | dmfs/defs | DM FT | DM FS | Method of caries identification | Clinical examination criteria | Outcome of interest: Fluorosis | Fluorosis (Dean's index) | Fluorosis (Thylstrup-Fejerskov index) | Tooth Surfaces Index of Fluorosis | Type of teeth examined for fluorosis | Hypomineralisation by photographs |
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Table 7 Caries outcome data form using example of primary dentition dmft

| Country | Author | Year | Age in years | CWF ppm | Baseline dmft CWF | Baseline CWF SD | Baseline CWF Total | Final dmft CWF | Final dmft SD CWF | Final CWF Total | Fluoride deficient ppm | Baseline mean dmft No F | Baseline SD No F | Baseline CWF Total | Final dmft No F | Final SD No F | Final No F Total | Difference in % point or dmft |
|---------|--------|------|--------------|---------|-------------------|-----------------|--------------------|----------------|-------------------|-----------------|------------------------|-------------------------|------------------|--------------------|-----------------|---------------|------------------|-------------------------------|
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This table was repeated for dmfs, % with CDC, and % without CDC for primary dentition. The table was also repeated for DMFT, DFMS, % with CDC, and % without CDC for permanent dentition

Table 8 Fluorosis outcome data form

| Country | Author | Year | Age in years | CWF ppm | Baseline % fluorosis | Baseline 95% CI | Baseline CWF affected number | Baseline CWF Total | Final % fluorosis | Final 95% CI | Final CWF affected number | Final CWF Total | Baseline 5 fluorosis No F | Baseline 95% CI No F | Baseline affected number No F | Baseline CWF Total | Final % No F | Final 95% CI No F | Final affected number No F | Final No F Total | Difference in % point or dmft |
|---------|--------|------|--------------|---------|----------------------|-----------------|------------------------------|--------------------|-------------------|--------------|---------------------------|-----------------|---------------------------|----------------------|-------------------------------|--------------------|--------------|-------------------|----------------------------|------------------|-------------------------------|
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6.5 Appendix E Quality assessment tool

Table 9 National Heart, Lung, and Blood Institute's (NHLBI's) quality assessment tool for observational cohort studies and cross-sectional surveys

| Number | NHLBI's quality assessment tool for observational cohort studies and cross-sectional surveys | Critical or non-critical |
|--------|--|--------------------------|
| 1 | <p>Was the research question or objective in this paper clearly stated (PECO)?</p> <p>Did the authors describe their research objective? Is it easy to understand what they were seeking to find? This issue is important for any scientific paper of any type. Higher quality scientific research explicitly defines a research question. (Population Exposure Comparator and Outcome for cohort and cross-sectional study designs, Population, Case, Controls and Exposure for case control, population, and outcome)</p> | |
| 2 | <p>Was the study population clearly specified and defined?</p> <p>Did the authors describe the group of people from which the study participants were selected or recruited, using demographics, location, and time-period? If you were to conduct this study again, would you know who to recruit, from where, and from what time-period? Is the cohort population free of the outcomes of interest at the time they were recruited?</p> <p>An example would be men over 40 years old with type 2 diabetes who began seeking medical care at Phoenix Good Samaritan Hospital between January 1, 1990 and December 31, 1994. In this example, the population is clearly described as: (1) who (men over 40 years old with type 2 diabetes); (2) where (Phoenix Good Samaritan Hospital); and (3) when (between January 1, 1990 and December 31, 1994). Another example is women ages 34 to 59 years of age in 1980 who were in the nursing profession and had no known coronary disease, stroke, cancer, hypercholesterolemia, or diabetes, and were recruited from the 11 most populous States, with contact information obtained from State nursing boards.</p> <p>In cohort studies, it is crucial that the population at baseline is free of the outcome of interest. For example, the nurses' population above would be an appropriate group in which to study incident coronary disease. This information is usually found either in descriptions of population recruitment, definitions of variables, or inclusion/exclusion criteria.</p> <p>You may need to look at prior papers on methods to make the assessment for this question. Those papers are usually in the reference list.</p> <p>For cross-sectional studies, a representative and adequately sized sample is required. The sample must be selected using randomisation or quasi randomised techniques (sequential or cluster sampling). A census of the population is also acceptable. The respondents must be similar to non-respondents, and this should be tested statistically.</p> <p>For case-control studies, a validated case (disease) definition is required, and the cases must meet the case definition and be representative of other cases; generally, community-based controls are considered the best comparators. The controls must have the same profile as the cases but without the disease of interest. The control number can exceed the case numbers by 4:1.</p> | |
| 3 | <p>Was the participation (response rate) of eligible persons at least 50%?</p> <p>If fewer than 50% of eligible persons participated in the study, then there is concern that the study population does not adequately represent the target population. This increases the risk of bias.</p> <p>This applies to three study designs.</p> | Critical flaw |
| 4 | <p>Were all the subjects selected or recruited from the same or similar populations (including the same time-period)? or Were inclusion and</p> | Critical flaw |

| Number | NHLBI's quality assessment tool for observational cohort studies and cross-sectional surveys | Critical or non-critical |
|--------|---|--------------------------|
| | <p>exclusion criteria for being in the study prespecified and applied uniformly to all participants?</p> <p>Were the inclusion and exclusion criteria developed prior to recruitment or selection of the study population? Were the same underlying criteria used for all the subjects involved? This issue is related to the description of the study population, above, and you may find the information for both questions in the same section of the paper.</p> <p>Most cohort studies begin with the selection of the cohort; participants in this cohort are then measured or evaluated to determine their exposure status. However, some cohort studies may recruit or select exposed participants in a different time or place than unexposed participants, especially retrospective cohort studies—which is when data are obtained from the past (retrospectively), but the analysis examines exposures prior to outcomes. For example, one research question could be whether diabetic men with clinical depression are at higher risk for cardiovascular disease than those without clinical depression. So, diabetic men with depression might be selected from a mental health clinic, while diabetic men without depression might be selected from an internal medicine or endocrinology clinic. This study recruit's groups from different clinic populations, so this example would get a 'no'.</p> <p>However, the women nurses described in the question above were selected based on the same inclusion/exclusion criteria, so that example would get a 'yes'.</p> <p>For cross-sectional studies, a representative and adequately sized sample is required. The sample must be selected using randomisation or quasi randomised techniques (sequential or cluster sampling). A census of the population is also acceptable. The respondents must be like non-respondents, and this should be tested statistically.</p> <p>For case-control studies, a validated case (disease) definition is required, and the cases must meet the case definition and be representative of other cases; generally, community-based controls are considered the best comparators. The controls must have the same profile as the cases but without the disease of interest. The control number can exceed the case numbers by 4:1.</p> | |
| 5 | <p>Was a sample size justification, power description, or variance (confidence intervals) and effect estimates (difference in effect between intervention and outcome) provided?</p> <p>*A confidence interval is the range of values (for example, proportions) in which the true value is likely to be found with a degree of certainty (by convention 95% degree of confidence), that is, the range of values will include the true value 95% of the time. It is an adjustment of the sample size calculation and demonstrates variance between the study sample and study population.</p> <p>Did the authors present their reasons for selecting or recruiting the number of people included or analysed (sample size calculator taking account of precision and effect size required sample size adjusted for response rate and loss to follow-up)? Do they note or discuss the statistical power of the study (effect size)? This question is about whether the study had enough participants to detect an association if one truly existed. This applies to cohort and cross-sectional studies.</p> <p>A paragraph in the methods section of the article may explain the sample size needed to detect a hypothesized difference in outcomes. You may also find a discussion of power in the discussion section (such as the study had 85% power to detect a 20% increase in the rate of an outcome of interest,</p> | Critical flaw |

| Number | NHLBI's quality assessment tool for observational cohort studies and cross-sectional surveys | Critical or non-critical |
|--------|---|--------------------------|
| | <p>with a 2-sided alpha of 0.05). Sometimes estimates of variance and/or estimates of effect size are given, instead of sample size calculations. In any of these cases, the answer would be 'yes'.</p> <p>However, observational cohort studies often do not report anything about power or sample sizes because the analyses are exploratory in nature. In this case, the answer would be 'no'. This is not a 'fatal flaw'. It just may indicate that attention was not paid to whether the study was sufficiently sized to answer a prespecified question—i.e., it may have been an exploratory, hypothesis-generating study.</p> | |
| 6 | <p>For the analyses of cohort studies, were the exposure(s) of interest measured prior to the outcome(s) being measured (temporal sequence, causality criteria)?</p> <p>This question is important because, to determine whether an exposure causes an outcome, the exposure must come before the outcome. For some prospective cohort studies, the investigator enrolls the cohort and then determines the exposure status of various members of the cohort (large epidemiological studies like Framingham used this approach). However, for other cohort studies, the cohort is selected based on its exposure status, as in the example above of depressed diabetic men (the exposure being depression). Other examples include a cohort identified by its exposure to fluoridated drinking water and then compared to a cohort living in an area without fluoridated water, or a cohort of military personnel exposed to combat in the Gulf War compared to a cohort of military personnel not deployed in a combat zone.</p> <p>With either of these types of cohort studies, the cohort is followed forward in time (i.e., prospectively) to assess the outcomes that occurred in the exposed members compared to nonexposed members of the cohort. Therefore, you begin the study in the present by looking at groups that were exposed (or not) to some biological or behavioural factor, intervention, etc., and then you follow them forward in time to examine outcomes. If a cohort study is conducted properly, the answer to this question should be 'yes', since the exposure status of members of the cohort was determined at the beginning of the study before the outcomes occurred.</p> <p>For retrospective cohort studies, the same principal applies. The difference is that, rather than identifying a cohort in the present and following them forward in time, the investigators go back in time (i.e., retrospectively) and select a cohort based on their exposure status in the past and then follow them forward to assess the outcomes that occurred in the exposed and nonexposed cohort members. Because in retrospective cohort studies the exposure and outcomes may have already occurred (it depends on how long they follow the cohort), it is important to make sure that the exposure preceded the outcome.</p> <p>Sometimes cross-sectional studies are conducted (or cross-sectional analyses of cohort-study data), where the exposures and outcomes are measured during the same timeframe. As a result, cross-sectional analyses provide weaker evidence than regular cohort studies regarding a potential causal relationship between exposures and outcomes. For cross-sectional analyses, the answer to Question 6 should be 'no'.</p> <p>For cross-sectional studies, the exposure status and outcome status are measured at the same time. Validated and in if possible objective measures should be used to assess the exposures and outcomes for all respondents. Attempts should be made to date the exposure and the outcome.</p> <p>For case-control studies, a validated case (disease) definition is required, and the cases must meet the case definition and be representative of other</p> | |

| Number | NHLBI's quality assessment tool for observational cohort studies and cross-sectional surveys | Critical or non-critical |
|--------|--|--------------------------|
| | cases; generally, community-based controls are considered the best comparators. The controls must have the same profile as the cases but without the disease of interest. The control number can exceed the case numbers by 4:1. Validated and in if possible objective measures should be used to assess the exposures and outcomes for cases and controls. In addition, the exposure status should be validated in existing records, dated, and ideally through objective testing. | |
| 7 | <p>Was the timeframe sufficient so that one could reasonably expect to see an association between exposure and outcome if it existed?</p> <p>Did the study allow enough time for sufficient outcomes to occur or be observed, or enough time for an exposure to have a biological effect on an outcome? In the examples given above, if clinical depression has a biological effect on increasing risk for cardiovascular disease, such an effect may take years. In the other example, if higher dietary sodium increases blood pressure, a short timeframe may be sufficient to assess its association with blood pressure, but a longer timeframe would be needed to examine its association with heart attacks.</p> <p>The issue of timeframe is important to enable meaningful analysis of the relationships between exposures and outcomes to be conducted. This often requires at least several years, especially when looking at health outcomes, but it depends on the research question and outcomes being examined. Cross-sectional analyses allow no time to see an effect, since the exposures and outcomes are assessed at the same time, so those would get a 'no' response.</p> <p>The time frame also applies to case-control studies.</p> | |
| 8 | <p>For exposures that can vary in amount or level, did the study examine different levels of the exposure as related to the outcome (e.g., categories of exposure, or exposure measured as continuous variable to determine dose response rate, causality criteria)?</p> <p>Levels of exposure applies to cross-sectional and cohort studies. If the exposure can be defined as a range (examples: drug dosage, amount of physical activity, amount of sodium consumed), were multiple categories of that exposure assessed? (for example, for drugs: not on the medication, on a low dose, medium dose, high dose; for dietary sodium, higher than average consumption in the United States of America, lower than recommended consumption, between the two). Sometimes discrete categories of exposure are not used, but instead exposures are measured as continuous variables (for example, mg/day of dietary sodium or blood pressure values).</p> <p>In any case, studying different levels of exposure (where possible) enables investigators to assess trends or dose-response relationships between exposures and outcomes—e.g., the higher the exposure, the greater the rate of the health outcome. The presence of trends or dose-response relationships lends credibility to the hypothesis of causality between exposure and outcome.</p> <p>For some exposures, however, this question may not be applicable (e.g., the exposure may be a dichotomous variable like living in a rural setting versus an urban setting, or vaccinated versus not vaccinated with a single dose and time vaccine). If there are only two possible exposures (yes/no), then this question should be given an 'not applicable' and it should not count negatively towards the quality rating.</p> | |
| 9 | Were the (different) exposure measures (independent variables that may cause the outcome) clearly defined, valid, reliable, and implemented consistently across all study participants? | |

| Number | NHLBI's quality assessment tool for observational cohort studies and cross-sectional surveys | Critical or non-critical |
|--------|--|--------------------------|
| | <p>Were the exposure measures defined in detail? Were the tools or methods used to measure exposure accurate and reliable—for example, have they been validated or are they objective? This issue is important as it influences confidence in the reported exposures. When exposures are measured with less accuracy or validity, it is harder to see an association between exposure and outcome even if one exists. Also as important is whether the exposures were assessed in the same manner within groups and between groups; if not, bias may result.</p> <p>For example, retrospective self-report of dietary salt intake is not as valid and reliable as prospectively using a standardized dietary log plus testing participants' urine for sodium content. Another example is measurement of blood pressure, where there may be quite a difference between usual care, where clinicians measure blood pressure however it is done in their practice setting (which can vary considerably), and use of trained blood pressure assessors using standardized equipment (e.g., the same blood pressure device which has been tested and calibrated) and a standardized protocol (e.g., patient is seated for 5 minutes with feet flat on the floor, blood pressure is taken twice in each arm, and all four measurements are averaged). In each of these cases, the former would get a 'no' and the latter a 'yes.'</p> <p>Here is a final example that illustrates the point about why it is important to assess exposures consistently across all groups: If people with higher blood pressure (exposed cohort) are seen by their providers more frequently than those without elevated blood pressure (nonexposed group), it also increases the chances of detecting and documenting changes in health outcomes, including cardiovascular disease-related events. Therefore, it may lead to the conclusion that higher blood pressure leads to more cardiovascular disease events. This may be true, but it could also be because the subjects with higher blood pressure were seen more often; thus, more cardiovascular disease-related events were detected and documented simply because they had more encounters with the health care system. Thus, it could bias the results and lead to an erroneous conclusion.</p> <p>Different exposure measures apply to both cross-sectional and case-control studies.</p> | |
| 10 | <p>Was the exposure(s) assessed more than once over time (causality, consistency)?</p> <p>Was the exposure for each person measured more than once during the study period? Multiple measurements with the same result increase our confidence that the exposure status was correctly classified. Also, multiple measurements enable investigators to look at changes in exposure over time, for example, people who ate high dietary sodium throughout the follow-up period, compared to those who started out high then reduced their intake, compared to those who ate low sodium throughout. Once again, this may not be applicable in all cases. In many older studies, exposure was measured only at baseline. However, multiple exposure measurements do result in a stronger study design.</p> | |
| 11 | <p>Were the (different) outcome measures (dependent variables such as a disease) clearly defined, valid, reliable, and implemented consistently across all study participants?</p> <p>Were the outcomes defined in detail? Were the tools or methods for measuring outcomes accurate and reliable, for example, have they been validated or are they objective (laboratory test)? This issue is important because it influences confidence in the validity of study results. Also</p> | |

| Number | NHLBI's quality assessment tool for observational cohort studies and cross-sectional surveys | Critical or non-critical |
|--------|--|--------------------------|
| | <p>important is whether the outcomes were assessed in the same manner within groups and between groups.</p> <p>An example of an outcome measure that is objective, accurate, and reliable is death—the outcome measured with more accuracy than any other. But even with a measure as objective as death, there can be differences in the accuracy and reliability of how death was assessed by the investigators. Did they base it on an autopsy report, death certificate, death registry, or report from a family member? Another example is a study of whether dietary fat intake is related to blood cholesterol level (cholesterol level being the outcome), and the cholesterol level is measured from fasting blood samples that are all sent to the same laboratory. These examples would get a 'yes'. An example of a 'no' would be self-report by subjects that they had a heart attack, or self-report of how much they weigh (if body weight is the outcome of interest).</p> <p>Like the example in Question 9, results may be biased if one group (e.g., people with high blood pressure) is seen more frequently than another group (people with normal blood pressure) because more frequent encounters with the health care system increases the chances of outcomes being detected and documented.</p> | |
| 12 | <p>Were the outcome assessors blinded to the exposure status of participants?</p> <p>Blinding means that outcome assessors did not know whether the participant was exposed or unexposed. It is also sometimes called 'masking'. The objective is to look for evidence in the article that the person(s) assessing the outcome(s) for the study (for example, examining medical records to determine the outcomes that occurred in the exposed and comparison groups) is masked to the exposure status of the participant. Sometimes the person measuring the exposure is the same person conducting the outcome assessment. In this case, the outcome assessor would most likely not be blinded to exposure status because they also took measurements of exposures. If so, make a note of that in the comments section.</p> <p>As you assess this criterion, think about whether it is likely that the person(s) doing the outcome assessment would know (or be able to figure out) the exposure status of the study participants. If the answer is no, then blinding is adequate. An example of adequate blinding of the outcome assessors is to create a separate committee, whose members were not involved in the care of the patient and had no information about the study participants' exposure status. The committee would then be provided with copies of participants' medical records, which had been stripped of any potential exposure information or personally identifiable information. The committee would then review the records for prespecified outcomes according to the study protocol. If blinding was not possible, which is sometimes the case, mark 'not applicable' and explain the potential for bias.</p> | |
| 13 | <p>Was loss to follow-up after baseline 20% or less?</p> <p>Higher overall follow-up rates are always better than lower follow-up rates, even though higher rates are expected in shorter studies, whereas lower overall follow-up rates are often seen in studies of longer duration. Usually, an acceptable overall participant follow-up rate is considered 80% or more of participants whose exposures were measured at baseline are follow-up at each data collection point. However, this is just a general guideline. For example, a 6-month cohort study examining the relationship between dietary sodium intake and blood pressure level may have over 90% follow-</p> | Critical flaw |

| Number | NHLBI's quality assessment tool for observational cohort studies and cross-sectional surveys | Critical or non-critical |
|--------|--|--------------------------|
|--------|--|--------------------------|

up, but a 20-year cohort study examining effects of sodium intake on stroke may have only a 65% follow-up rate.

Not applicable to case-control or cross-sectional studies

| | | |
|----|---|---------------|
| 14 | Were key potential confounding variables considered in the design (restriction or matching), measured during the study, and adjusted statistically for their impact on the relationship between exposure(s) and outcome(s) (by either stratification or regression analysis) | Critical flaw |
|----|---|---------------|

***Confounding** is when a factor has an association with the exposure and can independently cause the outcome or disease. It can over or underestimate an effect of interest or association. A **confounding variable** (also confounding factor or confounder) is a variable that has a relationship with both the exposure and outcome variable.

Were key potential confounding variables measured and adjusted for, such as by statistical adjustment for baseline differences? Logistic regression or other regression methods are often used to account for the influence of variables not of interest.

This is a key issue in cohort studies because statistical analyses need to control for potential confounders, in contrast to a randomised controlled trial, where the randomisation process controls for potential confounders. All key factors that may be associated both with the exposure of interest and the outcome—that are not of interest to the research question—should be controlled for in the analyses.

For example, in a study of the relationship between cardiorespiratory fitness and cardiovascular disease events (heart attacks and strokes), the study should control for age, blood pressure, blood cholesterol, and body weight, because all these factors are associated both with low fitness and with cardiovascular disease events. Well-done cohort studies control for multiple potential confounders.

For cohort studies, restriction, stratification, and regression are appropriate to control for confounding and these studies can calculate incidence.

For case-control studies, matching, restriction, and conditional logistic regression are appropriate to control for confounding and odds ratios should be employed as these studies cannot calculate incidence.

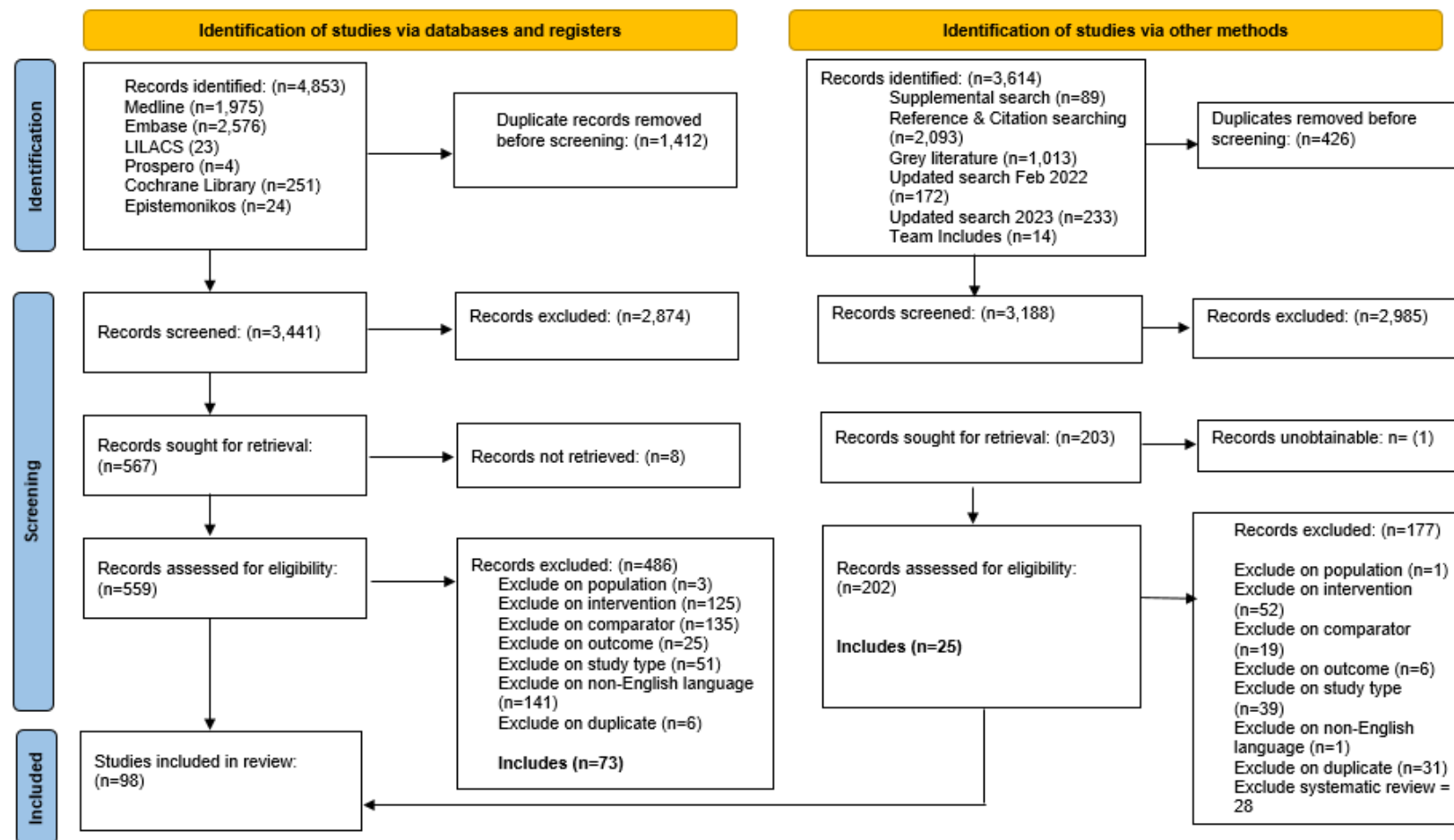
For cross-sectional studies, restriction, stratification, and regression are appropriate to control for confounding and proportions and odds ratios should be employed as these studies cannot calculate incidence.

Five questions were given more weight than the other questions; A negative scoring for one of these questions could be considered one critical flaw and a study with 3 or more critical flaws could be considered to contain evidence that the reader cannot trust:

1. Was the participation rate of eligible persons at least 50% (Question 3)?
2. Were all subjects selected or recruited from the same or similar populations (including same time period)? Were inclusion and exclusion criteria for being in the study prespecified and applied uniformly to all participants (Question 4)?
3. Was a sample size justification, power description, or variance and effect estimates provided (Question 5)?
4. Was loss to follow-up 20% or less (Question 13) (cohort only)?
5. Were key potential confounding variables measured and adjusted statistically for their impact on the relationship between exposure(s) and outcome(s) (Question 14)?

6.6 Appendix F PRISMA flow diagram for Question 1

PRISMA 2020 flow diagram for new systematic reviews which included searches of databases, registers, and other sources



From: Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. *BMJ* 2021;372:n71. doi: 10.1136/bmj.n71. For more information, visit: <http://www.prisma-statement.org/>

6.7 Appendix G Periodontal health results

We identified seven papers [117,127, 144-147,153] that examined periodontal disease published between 1972 and 1996. The studies were cross-sectional surveys. Three papers used Löe and Silness Gingival Index to assess aspects of periodontal health, two papers employed the Gingival Bleeding (sulcus) Index, one paper employed the periodontal index, and one did not report use of an index but presented the mean percentage of 6 index teeth with gingival bleeding.

- The Löe and Silness Gingival Index may be used for the assessment of prevalence and severity of gingivitis in populations, groups, and individuals. A score of zero indicates no inflammation. Scores of 0.1–1.0 signify mild inflammation, 1.1–2.0 signify moderate inflammation, and 2.1–3.0 signify severe inflammation.
- The Gingival Bleeding Index may be used for the assessment of prevalence and severity of gingivitis. There are two scores, 0 or 1, depending on whether or not bleeding occurs after a probe is gently run around the gingival sulcus. A percentage score is obtained by dividing by the number of teeth examined and multiplying the result by 100.
- The Periodontal Disease Index was created in 1978 by the World Health Organization to provide a global standard for screening periodontal disease in populations. The total of the scores for each tooth divided by the number of teeth examined: the higher the score, the more severe the periodontal disease.

In 1988, Clovis *et al.* reported that when all children were considered, regardless of length of residency in either non-fluoridated or fluoridated communities, there were no significant difference between the non-fluoridated and fluoridated communities in the severity of gingivitis using the Loe and Silness Gingival Index (mean index score \pm standard deviation for gingivitis in the CWF area was 0.58 ± 0.28 , $n=89$, compared with the mean index score in fluoride deficient area which was 0.69 ± 0.34 , $n=115$) [117]. The study was judged low quality with respect to design and implementation.

Hsieh *et al.* reported that the prevalence of gingivitis was very high among 6–12-year-old children living in two villages in Taiwan in 1971/2 prior to the introduction of CWF and the percentage range varied by age 96.4%–99.2%, the average periodontal index score was 0.4–0.5, and the average gingival unit with inflammation was 7.9–11.0 [127]. The study was judged moderate quality with respect to design and implementation.

Rugg-Gunn *et al.*, in 1977, reported that gingival inflammation in children living in the fluoride deficient area was more severe than in children living in CWF areas in both urban ($p<0.01$) and rural communities ($p<0.001$) using the Löe and Silness Gingival Index [147]. The mean gingival inflammation scores were higher in socially deprived urban areas with CWF at 0.85 than in the more affluent urban areas with CWF at 0.8 and rural CWF areas at 0.63. The mean gingival inflammation scores were also higher in more affluent urban and rural areas without CWF at 0.92 and 0.96, respectively, than in the socially deprived urban areas with CWF. The study was judged low quality with respect to design and implementation.

Rugg-Gunn *et al.*, in 1988, reported that gingival inflammation in children living in the nonfluoridated area was more severe than in children in the fluoridated area [146]. The earlier of the two studies use the same intervention areas but different comparator areas. The mean Löe and Silness Gingival Index scores were: fluoridated area 0.45 and non-fluoridated area 0.78 ($p < 0.001$). For social class III (more deprived) children only, the corresponding figures were 0.49 and 0.78 ($p < 0.001$). Of note, the gingival inflammation scores in the CWF area had fallen over the intervening 11 years, but according to the study authors the

decrease may reflect increased use of fluoridated toothpaste alongside CWF. The study was judged moderate quality with respect to design and implementation.

Parviainen *et al.* reported that the mean Gingival Bleeding Index score for the total sample in their study was 40% in 1973 [145]. The boys had more gingivitis than the girls ($p < 0.001$) and the mean Gingival Bleeding Index scores were 44% and 36%, respectively. No statistically significant linear shift in the Gingival Bleeding Index scores was observed with advancing age. There was more gingivitis in Hamina (2.5 ppm) than in Jyväskylä (0.2 ppm) or Kuopio (CWF 1.0 ppm). Parviainen *et al.*, in a follow-up survey in 1982, reported that the overall mean Gingival Bleeding Index score for the total study population in 1982 was 32% (compared with 40% in 1973). The bulk of this improvement was due to the improved gingival health among the boys. The significant difference in 1973 between total mean Gingival Bleeding Index score for boys and girls had disappeared in 1982 [144]. The decrease in the scores in the high-fluoride area (2.5 ppm) to 31% (compared with 50% in 1973) was highly significant ($p < 0.001$). A corresponding improvement was also observed in the optimal fluoride area (1.0 ppm), whereas in the low-fluoride area (0.2 ppm), the gingival conditions remained practically the same from 1973 to 1982. Because of these changes, the highly significant difference between the high-fluoride area and the two other areas in 1973 had disappeared in 1982. The study was judged low quality with respect to design and implementation.

Seppä *et al.* (1996) examined 12-year-old children for gingival bleeding after gentle probing of 6 index teeth (16, 11, 26, 36, 41 and 46) prior to their caries examination at baseline [153]. For each tooth, bleeding was considered present if found on any tooth surface. The results are presented as mean percentages of index teeth with gingival bleeding. Percentages of teeth with gingival bleeding were statistically significantly higher at 55% ($\pm 24\%$) in the CWF area compared with 39% ($\pm 24\%$) in the nonfluoridated area, $p < 0.0001$. The study design was a cross-sectional survey, and the quality assessment judged it to be low quality with respect to design and implementation.

6.8 Appendix H Complete quality assessment scores

Table 10 National Heart, Lung, and Blood Institute's (NHLBI's) quality assessment scores for observational cohort and cross-sectional studies for Question 1

| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6. | 7 | 8 | 9 | 10 | 11 Caries | 11 Fluorosis | 12 | 13 | 14 |
|-------------------|------|-----------|------------------------|-----|-----|------------------|-----|--------------------------|------------------|------------------|-----|-----|------------------|----------------|----------------|--------------|----------------|-----------|
| Medcalf | 1975 | Australia | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Not reported | Not applicable | Some |
| Carr | 1976 | Australia | Cross-sectional survey | Yes | Yes | Cannot determine | No | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Not applicable | Some |
| Riordan | 1991 | Australia | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | No | Not applicable | Partial |
| Riordan and Banks | 1991 | Australia | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | Not reported | Not applicable | Partial |
| Cortes et al. | 1996 | Brazil | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Cannot determine | Yes | Yes | Cannot determine | No | Yes | Partial | Not applicable | Some |
| Heintze et al. | 1998 | Brazil | Cross-sectional survey | Yes | Yes | No | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | Not reported | Not applicable | Some |
| Tiano et al. | 2009 | Brazil | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Cannot determine | Cannot determine | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Extensive |
| Tiano et al. | 2009 | Brazil | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Cannot determine | Cannot determine | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Extensive |
| Silva et al. | 2021 | Brazil | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | No | Yes | Not reported | Not applicable | Extensive |
| Brown | 1951 | Canada | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | No | No | Not reported | Not applicable | Some |
| Connor | 1963 | Canada | Cross-sectional survey | No | Yes | Yes | No | Not applicable as census | Yes | Yes | Yes | Yes | Cannot determine | No | No | Not reported | Not applicable | Some |
| Clovis et al. | 1988 | Canada | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Ismail et al. | 1990 | Canada | Cross-sectional survey | No | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Partial | Not applicable | Partial |
| Ismail et al. | 1993 | Canada | Cross-sectional survey | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Not reported | Not applicable | Some |

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| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6. | 7 | 8 | 9 | 10 | 11 Caries | 11 Fluorosi s | 12 | 13 | 14 |
|--------------------|-------|-------------|--|-----|-----|------------------|-----|----------------|------------------|------------------|-----|--------------|------------------|----------------|------------------|--------------|----------------|-----------|
| Clark et al. | 1994 | Canada | Cross-sectional survey | No | Yes | No | Yes | No | Yes | Cannot determine | Yes | Yes | Cannot determine | Not applicable | Yes | Not reported | Not applicable | Some |
| Maupomé et al. | 2003 | Canada | Cross-sectional survey | Yes | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | No | Not applicable | Partial? |
| Clark et al. | 2006 | Canada | Cross-sectional survey | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | Not reported | Not applicable | Some |
| McLaren et al. | 2017 | Canada | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Partial | Not applicable | Some |
| Maupomé et al. | 2001 | Canada | Retrospective/prospective cohort study | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | No | Partial |
| Brown et al. | 1960 | Canada | Cross-sectional survey | No | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | No | No | Not reported | Not applicable | Some |
| Brown and Poplove | 1965 | Canada | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | No | No | Not reported | Not applicable | Some |
| Clark et al. | 1993 | Canada | Cross-sectional survey | Yes | Yes | No | Yes | No | Cannot determine | Cannot determine | Yes | Yes | Cannot determine | Not applicable | Yes | Not reported | Not applicable | Some |
| Clark et al. | 1995 | Canada | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Cannot determine | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| McLaren et al. | 2021 | Canada | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Partial | Not applicable | Extensive |
| Villa et al. | 1998 | Chile | Cross-sectional survey | No | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Yes | Not applicable | Partial |
| Kunzel | 1982 | Cuba | Cross-sectional survey | No | Yes | Not reported | Yes | No | Cannot determine | Yes | Yes | Yes | Yes | Yes | Yes | No | Not applicable | Some |
| Kunzel and Fischer | 2000 | Cuba | Cross-sectional survey | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Beal and James | 1971 | England, UK | Cross-sectional survey | No | Yes | Not reported | Yes | No | Cannot determine | No | Yes | Not reported | Not reported | Yes | Not applicable | No | Not applicable | Some |
| Jackson et al. | 1975b | England | Cross-sectional survey | Yes | Yes | Yes | Yes | Not applicable | yes | Yes | Yes | Not reported | Not reported | Yes | Not applicable | Yes | Not applicable | Some |

HRB Document Template

| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6. | 7 | 8 | 9 | 10 | 11 Caries | 11 Fluorosi s | 12 | 13 | 14 |
|--------------------|------|-------------|--|-----|-----|--------------|-----|----------------|-----|-----|-----|--------------|--------------|----------------|------------------|-----|----------------|------|
| Rugg-Gunn et al. | 1977 | England, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Jackson et al. | 1980 | England, UK | Cross-sectional survey | Yes | Yes | Not reported | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Beal and Clayton | 1981 | England, UK | Cross-sectional survey | Yes | Yes | Not reported | Yes | No | No | No | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Rugg-Gunn et al. | 1981 | England, UK | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Hardwick et al | 1982 | England, UK | Retrospective/prospective cohort study as 4 year follow-up for the same children | Yes | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | Yes | No | Some |
| French et al. | 1984 | England, UK | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Mitropoulos et al. | 1988 | England, UK | Cross-sectional survey | Yes | Yes | Not reported | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Rugg-Gunn et al. | 1988 | England, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Murray et al. | 1991 | England, UK | Cross-sectional survey | Yes | Yes | Not reported | Yes | No | Yes | Yes | Yes | Not reported | Not reported | Yes | Not applicable | No | Not applicable | Some |
| Booth et al. | 1992 | England, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Evans et al. | 1995 | England, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Evans et al. | 1996 | England, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Tabari et al. | 2000 | England, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | Yes | Yes | Not applicable | Some |

HRB Document Template

| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6. | 7 | 8 | 9 | 10 | 11 Caries | 11 Fluorosi s | 12 | 13 | 14 |
|------------------------|------|--------------------|--|-----|-----|------------------|-----|-----|-----|-----|-----|--------------|------------------|--------------|---------------------|--------------|----------------|---------|
| Gray and Davies-Slowik | 2001 | England, UK | Cross-sectional survey | No | Yes | Not reported | Yes | No | Yes | Yes | Yes | Not reported | Not reported | Yes | Not applicable | No | Not applicable | Some |
| Goodwin et al. | 2022 | England, UK | Retrospective/prospective cohort study | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | No | Some |
| Ellwood and O'Mullane | 1995 | England, Wales, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Ellwood and O'Mullane | 1996 | England, Wales, UK | Cross-sectional survey | No | Yes | Not reported | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | None |
| Parviainen et al. | 1977 | Finland | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Hausen et al. | 1981 | Finland | Retrospective/prospective cohort study | Yes | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Not applicable | Partial |
| Parviainen et al. | 1985 | Finland | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Partial |
| Linkosalo | 1986 | Finland | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Not applicable | Some |
| Seppa et al. | 1996 | Finland | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Partial |
| Seppa et al. | 1998 | Finland | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Partial |
| Seppa et al. | 2000 | Finland | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | No | Not applicable | Partial |
| Seppa et al. | 2000 | Finland | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | No | Not applicable | Partial |
| Seppa et al. | 2002 | Finland | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Not applicable | Some |

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| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6. | 7 | 8 | 9 | 10 | 11 Caries | 11 Fluorosi s | 12 | 13 | 14 |
|------------------------|------|-------------|-------------------------------|-----|-----|------------------|-----|-----|------------------|--|-----|-----|------------------|----------------|---------------------|--------------|----------------|---------|
| Kunzel | 1968 | German y | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Not applicable | Some |
| Kunzel | 1980 | German y | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Not applicable | Some |
| Kunzel and Fischer | 1997 | German y | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Kunzel et al. | 2000 | German y | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Lemasney et al. | 1984 | Ireland | Cross-sectional survey | No | Yes | Not reported | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| O'Mullane et al. | 1986 | Ireland | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not reported | Not applicable | Some |
| O'Mullane et al. | 1988 | Ireland | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Cannot determine | sources to the public principally through fluoridation of water supplies | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| Clarkson and O'Mullane | 1992 | Ireland | Cross-sectional survey | No | Yes | Not reported | Yes | No | Yes | Yes | Yes | Yes | Yes | Not applicable | Yes | | Not applicable | Some |
| Whelton et al. | 2004 | Ireland | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | No | Not applicable | Some |
| Mullen et al. | 2012 | Ireland | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Some |
| James et al. | 2021 | Ireland | Cross-sectional survey/cohort | No | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | No | Yes | Partial |
| Mohd Nor et al. | 2018 | Malaysia | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Partial | Not applicable | Partial |

HRB Document Template

| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6. | 7 | 8 | 9 | 10 | 11 Caries | 11 Fluorosi s | 12 | 13 | 14 |
|------------------------|------|--------------|--|-----|-----|------------------|-----|----------------|------------------|------------------|----------------|------------------|------------------|----------------|------------------|--------------|------------------|---------|
| Mohd Nor et al. | 2021 | Malaysia | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | Partial | Not applicable | Partial |
| Backer Dirks et al. | 1961 | Netherlands | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Yes | Not applicable | Some |
| Groeneveld | 1985 | Netherlands | Retrospective/prospective cohort study | Yes | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Cannot determine | Some |
| Kalsbeek et al. | 1993 | Netherlands | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Not applicable | Some |
| Weerheijm et al. | 1997 | Netherlands | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| de Liefde and Herbison | 1985 | New Zealand | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Not reported | Not applicable | Some |
| Treasure and Dever | 1992 | New Zealand | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Treasure and Dever | 1994 | New Zealand | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Cannot determine | Cannot determine | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Ministry of Health | 2010 | New Zealand | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Not reported | Not applicable | Some |
| Stephen et al. | 1987 | Scotland, UK | Cross-sectional survey | Yes | Yes | Not reported | No | Not applicable | Yes | Yes | Yes | Cannot determine | Not reported | Yes | Not applicable | No | Not applicable | None |
| Wong et al. | 1970 | Singapore | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | No | Not reported | Not applicable | Some |
| Hsieh et al. | 1972 | Taiwan | Cross-sectional survey | Yes | Yes | Yes | Yes | Not applicable | Not applicable | Not applicable | Not applicable | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Hsieh et al. | 1979 | Taiwan | Cross-sectional survey | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Guo et al. | 1984 | Taiwan | Cross-sectional survey | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |

HRB Document Template

| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6. | 7 | 8 | 9 | 10 | 11 Caries | 11 Fluorosis | 12 | 13 | 14 |
|------------------|-------|-----------|-------------------------------|-----|-----|------------------|------------------|----------------|-----|-----|-----|-----|------------------|----------------|----------------|--------------|------------------|-----------|
| Hsieh et al. | 1986 | Taiwan | Cross-sectional survey | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Hong et al. | 1990 | Taiwan | Cross-sectional survey | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | Not reported | Not applicable | Some |
| Ast et al. | 1951 | USA | Cross-sectional survey series | Yes | Yes | Cannot determine | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Partial | Not applicable | Some |
| Ast and Chase | 1953 | USA | Cross-sectional survey | No | Yes | Cannot determine | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Cannot determine | Some |
| Szpunar and Burt | 1988 | USA | Cross-sectional survey | Yes | Yes | No | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Yes | No | Not applicable | Some |
| Gillcrist et al. | 2001 | USA | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | Not reported | Not applicable | Some |
| Ast et al. | 1950 | USA | Cross-sectional survey | Yes | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Cannot determine | Some |
| Arnold et al. | 1953 | USA | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Not applicable | Some |
| Ast et al. | 1955 | USA | Cross-sectional survey | No | Yes | Cannot determine | Cannot determine | No | Yes | Yes | Yes | Yes | Cannot determine | No | Not applicable | Not reported | Not applicable | Some |
| Arnold et al. | 1956 | USA | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | No | Yes | Not reported | Not applicable | Some |
| Kumar et al. | 1989 | USA | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Partial | Not applicable | Some |
| Kumar et al. | 1998 | USA | Cross-sectional survey | No | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Not reported | Not applicable | Some |
| Kumar et al. | 2000 | USA | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | | Not applicable | Extensive |
| Jackson et al. | 1975a | Wales, UK | Cross-sectional survey | Yes | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Yes | Not applicable | None |
| Jackson et al. | 1985 | Wales, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Yes | Not applicable | None |

| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6. | 7 | 8 | 9 | 10 | 11 Caries | 11 Fluorosi s | 12 | 13 | 14 |
|---------------|------|-----------|------------------------|-----|-----|-----|-----|----|-----|-----|-----|-----|------------------|--------------|---------------------|----|----------------|------|
| Seaman et al. | 1989 | Wales, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Cannot determine | Yes | Not applicable | No | Not applicable | None |

*1. Research question stated

2. Study population clearly specified

3. Participation rate at least 50%

4. Subjects selected from the same population and inclusion and exclusion criteria prespecified

5. Sample size justification, power description, or variance and effect estimates provided

6. Exposure(s) of interest measured prior to outcome(s) measure

7. Timeframe sufficient to see an association between exposure and outcome

8. For exposures, study examine different levels of the exposure as related to the outcome

9. Exposure measures defined, valid, reliable, and consistently applied

10. Exposure(s) assessed more than once

11. Outcome measures defined, valid, reliable, and consistently applied: Caries

11. Outcome measures defined, valid, reliable, and consistently applied: Fluorosis

12. Outcome assessors blinded to the exposure status

13. Loss to follow-up 20% or less

14. Potential confounding exposures measured and adjusted statistically in outcomes

For each paper reporting on a longitudinal cohort study, cross-sectional survey, or case-control study, the scores were summed (for a total score ranging from 0.0 to 5.0). Papers scoring less than 3.0 were rated 'low quality', papers scoring 3.0 were rated 'moderate quality', and papers scoring 3.5 or more were rated 'high quality'. As many studies were cross-sectional in nature (point-in-time surveys) and scored 0.0 on item 13 (loss to follow-up not applicable), the maximum possible score for papers reporting on these types of studies was effectively capped at 4.0; for this reason, the threshold for 'high quality' was set at 3.5, rather than 4.0, in order to allow more effective differentiation of papers at the upper end of the range of scores. We also report the quality deficiencies by low-, moderate- and high-quality papers.

6.9 Appendix I Feasibility assessment results

Table 11 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF on dental caries, dmft

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure and proportion agreement where reported | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|------------------------------------|---|------------------------|-----------------------------|------------------------------------|---|----------------------------------|--------------------------------------|---|---------------|---------------------------------------|
| Silva et al. 2021, Brazil | Cross-sectional survey | 5 | 0.5 to 0.6 ppm | <0.05 ppm | dmft (0.92) | Mean and SD | Logistic regression analysis | Age, gender, lifetime exposure, socio-economic status, mother's level of education Snacks (sugar ingestion), CWF, fluoride toothpaste and toothbrushing Assess visits to dentist. | High | Yes |
| Tiano et al. 2009a, Brazil | Cross-sectional survey | 1-2 | 0.60–0.75ppm | <0.40 ppm | dmft | Mean and SD | No | Not applicable | Moderate | No, exclude on age, below 5 years old |
| McLaren et al. 2021, Canada | Cross-sectional survey and adjusted for cluster sampling but design effect not reported | 7 | 0.6 to 0.8 ppm | 0.07–0.30 ppm | dmft (≥ 0.80 most of the time) | Mean and SD | Poisson and logistic regression | Child's oral health, tooth brushing twice/day, floss once/day, visit dentist, last time visit dentist, dental insurance, fruit and vegetables at least once/day, Sugary drinks, fluoride supplements at home, fluoride treatments at the dentist's office, fluoride treatments in school programme, use of fluoride toothpaste, and/or fluoride mouth wash, household education, home ownership, and ethnocultural background | High | Yes |
| Villa et al. 1998, Chile | Cross-sectional survey and adjusted for cluster sampling but design effect not reported | 7 | 0.93ppm | 0.07 ppm | dmft | Mean and SD | No | Age, social economic status | Moderate | Yes |
| Jackson et al. 1975b, England UK | Cross-sectional survey | 5 | 1.0 ppm | <0.1 ppm | dmft | Mean and SD | No | Not applicable | Low | Yes |
| Rugg-Gunn et al. 1977, England, UK | Cross-sectional survey | 5 | 1.0ppm | <0.1 ppm | dmft | Mean and SD | No | Not applicable | Low | Yes |
| Jackson et al. 1980, England, UK | Cross-sectional survey | 5 | 0.9ppm | <0.1 ppm | dmft | Mean and SD | No | Not applicable | Low | Yes |
| Beal and Clayton 1981, England, UK | Cross-sectional survey | 5 | 0.85–0.90ppm | <0.35 ppm | dmft | Mean and SD | No | Not applicable | Low | Yes |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure and proportion agreement where reported | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--------------------------------------|---|------------------------|-----------------------------|------------------------------------|---|---------------------------------------|--------------------------------------|--|---------------|-----------------------------------|
| Rugg-Gunn et al. 1981, England, UK | Cross-sectional survey | 5 | 1.0ppm | <0.1ppm | dmft | Mean and SD | No | Not applicable | Low | Yes |
| French et al. 1984, England, UK | Cross-sectional survey | 5 | 1.0ppm | 0.1 ppm | dmft | Mean and SD | No | Not applicable | Low | Yes |
| Rugg-Gunn et al. 1988, England, UK | Cross-sectional survey | 5 | 1.0ppm | <0.1ppm | dmft | Mean and SD | No | Not applicable | Moderate | Yes |
| Booth et al. 1992, England, UK | Cross-sectional survey | 3 | 1.0ppm | <0.1ppm | dmft | Mean and SD | No | Not applicable | Moderate | No, exclude on age, below 5 years |
| Evans et al. 1995, England, UK | Cross-sectional survey | 5 | 1.0 ppm | < 0.1 ppm | dmft | Mean, variance not required as census | No | Not applicable | Moderate | Yes |
| Goodwin et al. 2022, England, UK | Retrospective/prospective cohort study | 5 | 0.9ppm | <0.2ppm | dmft (0.75 - 1.0) | Mean and SD | Negative binomial regression | Age, sex, and deprivation | Moderate | Yes |
| Lemasney et al. 1984, Ireland | Cross-sectional survey | 5 | 0.8–1.0 ppm | ≤3 ppm | dmft (Inter 0.96 - 0.98 Intra 0.98 - 1.0) | Mean and SD | No | Not applicable | Low | Yes |
| O'Mullane et al. 1986, Ireland | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 5 | 0.8–1.0 ppm | ≤3 ppm | dmft (>0.95 correlation coefficients) | Mean and SD | No | Not applicable | Moderate | Yes |
| Whelton et al. 2004, Ireland | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 5 | 0.8–1.0 ppm | ≤3 ppm | dmft | Mean and SD | No | Not applicable | Moderate | Yes |
| James et al. 2021, Ireland | Cross-sectional survey | 8 | 0.8–1.0 ppm | ≤3 ppm | dmft (Intra 0.86 to 1.00 in 2002, 0.77 to 1.00 in 2017) | Mean and SD | Multivariate regression | age, gender, age first used toothpaste, amount of toothpaste, frequency of toothbrushing, age first visited dentist, rinse method after toothbrushing, and sweet snacks between meals. | High | Yes |
| Treasure and Dever 1992, New Zealand | Cross-sectional survey, description indicate that authors have adjusted | 5 | 1.0 ppm | 0.08 ppm | dmft | Mean and SD | No | Not applicable | Moderate | Yes |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure and proportion agreement where reported | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--------------------------------------|---|------------------------|-----------------------------|------------------------------------|---|---------------------------------------|--------------------------------------|----------------|---------------|---|
| | for cluster sampling but not reported | | | | | | | | | |
| Ministry of Health 2010, New Zealand | Cross-sectional survey and adjusted for cluster sampling using a design effect of ≥ 2 | 5 | 0.8–0.9 ppm | 0.15 ppm | dmft (≥ 0.78 ICC) | Mean and SD | No | Not applicable | Moderate | No, exclude on population, numbers not reported |
| Guo et al. 1984, Taiwan | Cross-sectional census survey | 5 | 0.6ppm | 0.08 ppm | dmft | Mean, variance not required as census | No | Not applicable | Moderate | Yes |
| Hsieh et al. 1986, Taiwan | Cross-sectional census survey | 5 | 0.6ppm | 0.08 ppm | dmft | Mean, variance not required as census | No | Not applicable | Moderate | Yes |
| Jackson et al. 1985, Wales, UK | Cross-sectional survey | 5 | 0.99 ppm | <0.1ppm | dmft | Mean and SD | No | Not applicable | Moderate | Yes |
| Seaman et al. 1989, Wales, UK | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 5 | 0.8 ppm | <0.1ppm | dmft (0.86) | Mean and SD | No | Not applicable | Low | Yes |

Table 12 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF on dental caries, dmfs

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure and proportion agreement where reported | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--------------------------------------|---|------------------------|-----------------------------|------------------------------------|--|---------------------------------------|--------------------------------------|----------------|---------------|----------------------------|
| Tiano et al. 2009a, Brazil | Cross-sectional survey | 8-36 months | 0.60–0.75ppm | 0.4 ppm | dmfs | Mean and SD | No | Not applicable | Moderate | Yes |
| Rugg-Gunn et al. 1977, England, UK | Cross-sectional survey | 5 | 1.0ppm | <0.1 ppm | dmfs | Mean and SD | No | Not applicable | Low | Yes |
| Rugg-Gunn et al. 1981, England, UK | Cross-sectional survey | 5 | 1.0ppm | <0.1 ppm | dmfs | Mean and SD | No | Not applicable | Low | Yes |
| French et al. 1984, England, UK | Cross-sectional survey | 5 | 1.0ppm | <0.1 ppm | dmfs | Mean and SD | No | Not applicable | Low | Yes |
| Rugg-Gunn et al. 1988, England, UK | Cross-sectional survey | 5 | 1.0ppm | <0.1 ppm | dmfs | Mean and SD | No | Not applicable | Moderate | Yes |
| Evans et al. 1995, England, UK | Cross-sectional survey | 5 | 1.0 ppm | < 0.1 ppm | dmfs | Mean, variance not required as census | No | Not applicable | Moderate | Yes |
| Seppa et al. 2000b, Finland | Cross-sectional survey | 6 | 1.0ppm | <0.1 ppm | dmfs + radiographs (Inter 0.86 - 0.94, Intra 0.88 - 0.91, different years) | Mean and SD | No | Not applicable | Low | Yes |
| Treasure and Dever 1992, New Zealand | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 5 | 1.0 ppm | Not reported | dmfs | Mean and SD | No | Not applicable | Moderate | Yes |

Table 13 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF on dental caries, per cent without cavitated dental caries in the primary dentition

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure and proportion agreement where reported | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--------------------------------------|---|------------------------|-----------------------------|------------------------------------|---|----------------------------------|--------------------------------------|----------------|---------------|----------------------------|
| Brown et al. 1960, Canada | Cross-sectional survey | 9-11 | 1.0–1.2ppm | Not fluoridated | % primary teeth without CDC | %, 95% CI | No | Not applicable | Moderate | Yes |
| Gray and Davies-Slowik 2001, England | Cross-sectional survey | 5 | 1.0ppm | <0.3 ppm | % primary teeth without CDC | %, 95% CI | No | Not applicable | Low | Yes |
| Gillcrist et al. 2001, USA | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 5-11 | 1.0ppm | <0.3 ppm | % primary teeth without CDC | %, 95% CI | No | Not applicable | Low | Yes |

Table 14 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF on dental caries, per cent with cavitated dental caries in the primary dentition

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure and proportion agreement where reported | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--------------------------------|---|------------------------|-----------------------------|------------------------------------|---|------------------------------------|--------------------------------------|---|---------------|----------------------------|
| McLaren et al. 2021, Canada | Cross-sectional survey and adjusted for cluster sampling but design effect not reported | ~7 | 0.6 to 0.8 | 0.07–0.30 ppm | % primary teeth with CDC (≥ 0.80 most of the time) | %, 95% CI | Poisson and logistic regression | Child's oral health, tooth brushing twice/day, floss once/day, visit dentist, last time visit dentist, dental insurance, fruit and vegetables at least once/day, Sugary drinks, fluoride supplements at home, fluoride treatments at the dentist's office, fluoride treatments in school programme, use of fluoride toothpaste, and/or fluoride mouth wash, household education, home ownership, and ethnocultural background | High | Yes |
| Evans et al. 1995, England, UK | Cross-sectional survey | 5 | 1.0 ppm | < 0.1 ppm | % primary teeth with CDC | %, variance not required as census | No | Not applicable | Moderate | Yes |
| Guo et al. 1984, Taiwan | Cross-sectional census survey | 5 | 0.6 ppm | 0.08 ppm | % primary teeth with CDC | %, variance not required as census | No | Not applicable | Moderate | Yes |
| Hsieh et al. 1986, Taiwan | Cross-sectional census survey | 5 | 0.6 -0.7 ppm | 0.08 ppm | % primary teeth with CDC | %, variance not required as census | No | Not applicable | Moderate | Yes |

Table 15 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF on dental caries, DMFT

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure (proportion agreement where reported) | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--------------------------------------|---|------------------------|-----------------------------|------------------------------------|---|----------------------------------|--------------------------------------|---|---------------|----------------------------|
| Silva et al. 2021, Brazil | Cross-sectional survey and adjusted for cluster sampling using a design effect of 1.7 | 12 | 0.5 to 0.6 ppm | <0.05 ppm | DMFT (0.92) | Mean and SD | Logistic regression analysis | Age, gender, lifetime exposure, socio-economic status, mother's level of education Snacks (sugar ingestion), CWF, fluoride toothpaste and toothbrushing Assess visits to dentist. | High | Yes |
| Clovis et al. 1988, Canada | Cross-sectional survey | 11-12 | 1.08 ppm | 0.23 ppm | DMFT | Mean and SD | No | Not applicable | Low | Yes |
| Brown and Poplove 1965, Canada | Cross-sectional survey | 16-17 | 1.0–1.2 ppm | NF | DMFT | Mean and SD | No | Not applicable | Low | Yes |
| Brown et al. 1960, Canada | Cross-sectional survey | 9-11 | 1.0–1.2ppm | NF | DMFT | Mean and SD | No | Not applicable | Moderate | Yes |
| McLaren et al. 2021, Canada | Cross-sectional survey and adjusted for cluster sampling but design effect not reported | 7 | 0.59–0.89 ppm | ≤3 ppm | DMFT (≥ 0.80 most of the time) | Mean and SD | Poisson and logistic regression | Child's oral health, tooth brushing twice/day, floss once/day, visit dentist, last time visit dentist, dental insurance, fruit and vegetables at least once/day, Sugary drinks, fluoride supplements at home, fluoride treatments at the dentist's office, fluoride treatments in school programme, use of fluoride toothpaste, and/or fluoride mouth wash, household education, home ownership, and ethnocultural background | High | Yes |
| Villa et al. 1998, Chile | Cross-sectional survey and adjusted for cluster sampling but design effect not reported | 12 | 0.93ppm | ≤3 ppm | DMFT (≥0.91) | Mean and SD | No | Age, socio-economic status | Moderate | Yes |
| Kunzel and Fischer 2000, Cuba | Cross-sectional survey | 10-11 | 0.8ppm | <0.3ppm | DMFT | Mean and SD | No | Not applicable | Moderate | Yes |
| Mitropoulos et al. 1988, England, UK | Cross-sectional survey | 14 | 1.0ppm | Not fluoridated | DMFT | Mean and SD | No | Not applicable | Low | Yes |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure (proportion agreement where reported) | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|-----------------------------------|---|------------------------|-----------------------------|------------------------------------|---|----------------------------------|--------------------------------------|----------------|---------------|----------------------------|
| Murray et al. 1991, England, UK | Cross-sectional survey | 15-16 | 1.0ppm | 0.07–0.30 ppm | DMFT | Mean and SD | No | Not applicable | Low | Yes |
| Kunzel 1980, Germany | Cross-sectional survey | 10 | 1.0ppm | 0.07 ppm | DMFT | Mean and SD | No | Not applicable | Low | Yes |
| Kunzel et al. 2000, Germany | Cross-sectional survey | 12 | 0.8–1.0ppm | 0.05-0.1 ppm | DMFT (Inter 0.95, Intra 0.89 - 92.7) | Mean and SD | No | Not applicable | Low | Yes |
| Lemasney et al. | Cross-sectional survey | 11 | 0.8–1.0 ppm | <0.1 ppm | DMFT (Inter 0.96 - 0.98 Intra 0.98 - 1.0) | Mean and SD | No | Not applicable | Low | Yes |
| O'Mullane et al. 1986, Ireland | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 12 | 0.8–1.0 ppm | 0 ppm | DMFT (>0.95 correlation coefficients) | Mean and SD | No | Not applicable | Moderate | Yes |
| Whelton et al. 2004, Ireland | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 12 | 0.8–1.0 ppm | 0.1ppm | DMFT | Mean and SD | No | Not applicable | Moderate | Yes |
| Mullen et al. 2012, Ireland | Cross-sectional survey | 16 | 0.7ppm | 0.2ppm | DMFT (>0.80) | Mean and SD | No | Not applicable | Moderate | Yes |
| Mohd Nor et al. 2018, Malaysia | Cross-sectional survey | 12 | 0.5ppm | Not fluoridated | DMFT | Mean and SD | No | Not applicable | Moderate | Yes |
| Kalsbeek et al. 1993, Netherlands | Cross-sectional survey | 15 | 1.1ppm | <0.1ppm | DMFT + radiographs (0.89,0.99,0.99 and 0.91 for DS, FS, DFS, | Mean and SD | No | Not applicable | Low | Yes |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure (proportion agreement where reported) | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--|---|------------------------|-----------------------------|------------------------------------|---|---------------------------------------|--------------------------------------|----------------|---------------|----------------------------|
| | | | | | total dental caries lesions, respectively) | | | | | |
| de Liefde and Herbison 1985, New Zealand | Cross-sectional survey | 9 | 1.0ppm | 0.2 ppm | DMFT | Mean and SD | No | Not applicable | Low | Yes |
| Treasure and Dever 1994, New Zealand | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 14 | 1.0ppm | 0.08 ppm | DMFT | Mean and SD | No | Not applicable | Moderate | Yes |
| Hsieh et al. 1979, Taiwan | Cross-sectional census survey | 6 | 0.6ppm and then 0.7ppm | 0.08 ppm | DMFT | Mean, variance not required as census | No | Not applicable | Moderate | Yes |
| Guo et al. 1984, Taiwan | Cross-sectional census survey | 10 | 0.6ppm | 0.08 ppm | DMFT | Mean, variance not required as census | No | Not applicable | Moderate | Yes |
| Hsieh et al. 1986, Taiwan | Cross-sectional census survey | 12 | 0.6ppm and then 0.7ppm | <0.1 ppm | DMFT | Mean, variance not required as census | No | Not applicable | Moderate | Yes |
| Jackson et al. 1975a, Wales, UK | Cross-sectional survey | 15 | 0.9ppm | 0.12 - 0.19 ppm | DMFT | Mean and SD | No | Not applicable | Low | Yes |
| Jackson et al. 1985, Wales, UK | Cross-sectional survey | 15 | 0.99ppm | <0.1 ppm | DMFT | Mean and SD | No | Not applicable | Moderate | Yes |
| Thomas and Kassab 1992, Wales, UK | Cross-sectional survey | 18-30 | 0.8ppm | <0.1 ppm | DMFT | Mean and SD | No | Not applicable | Moderate | Yes |

Table 16 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF on dental caries, DMFS

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure and proportion agreement where reported | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--|---|------------------------|-----------------------------|------------------------------------|---|----------------------------------|--------------------------------------|----------------|---------------|----------------------------|
| Kunzel and Fischer 2000, Cuba | Cross-sectional survey | 10-11 | 0.8 ppm | 0.05-0.1 ppm | DMFS | Mean and SD | No | Not applicable | Moderate | Yes |
| Murray et al. 1991, England, UK | Cross-sectional survey | 15-16 | 1.0 ppm | <0.1 ppm | DMFS | Mean and SD | No | Not applicable | Low | Yes |
| Ellwood and O'Mullane 1995, England, Wales, UK | Cross-sectional survey | 14 | 0.7 ppm | <0.1ppm | DMFS (>0.81) | Mean and SD | Multiple linear regression | SES | Low | Yes |
| Kalsbeek et al. 1993, Netherlands | Cross-sectional survey | 15 | 1.1 ppm | 0.1 ppm | DMFS + radiographs (0.89,0.99,0.99 and 0.91 for DS, FS, DFS, total dental caries lesions, respectively) | Mean and SD | No | Not applicable | Low | Yes |
| Treasure and Dever 1994, New Zealand | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 14 | 1.0 ppm | 0.08 ppm | DMFS | Mean and SD | No | Not applicable | Moderate | Yes |
| Gillcrist et al. 2001, USA | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 5-11 | 1.0 ppm | <0.3 ppm | DMFS | Mean and SD | No | Not applicable | Low | Yes |

Table 17 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF on dental caries, per cent without cavitated dental caries in the permanent teeth

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure and proportion agreement where reported | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--------------------------------|---|------------------------|-----------------------------|------------------------------------|---|----------------------------------|--------------------------------------|----------------|---------------|----------------------------|
| Brown et al. 1960, Canada | Cross-sectional survey | 12–14 | 1.0–1.2 ppm | Not fluoridated | % permanent teeth without CDC | %, 95% CI | No | Not applicable | Moderate | Yes |
| Brown and Poplove 1965, Canada | Cross-sectional survey | 16–17 | 1.0–1.2 ppm | Not fluoridated | % permanent teeth without CDC | %, 95% CI | No | Not applicable | Low | Yes |
| Gillcrist et al. 2001, USA | Cross-sectional survey, description indicate that authors have adjusted for cluster sampling but not reported | 5–11 | 1.0 ppm | <0.3 ppm | % permanent teeth without CDC | %, 95% CI | No | Not applicable | Low | Yes |

Table 18 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF on dental caries, per cent with cavitated dental caries in the permanent teeth

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population (age) | CWF lifetime exposure (ppm) | Comparator lifetime exposure (ppm) | Dental caries outcome measure and proportion agreement where reported | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|-----------------------------|---|------------------------|-----------------------------|------------------------------------|---|------------------------------------|--------------------------------------|--|---------------|----------------------------|
| McLaren et al. 2021, Canada | Cross-sectional survey and adjusted for cluster sampling but design effect not reported | 7 | 0.61–0.82 ppm | 0.07–0.30 ppm | % with CDC (≥ 0.80 most of the time) | %, 95% CI | Poisson and logistic regression | Child's oral health, tooth brushing twice/day, floss once/day, visit dentist, last time visit dentist, dental insurance, fruits and vegetables at least once/day, Sugary drinks, fluoride supplements at home, fluoride treatments at the dentist's office, fluoride treatments in school programme, use of fluoride toothpaste, and/or fluoride mouth wash, household education, home ownership, and ethnocultural background | High | Yes |
| Guo et al. 1984, Taiwan | Cross-sectional census survey | 10 | 0.6 ppm | 0.08 ppm | % with CDC | %, variance not required as census | No | Not applicable | Moderate | Yes |
| Hsieh et al. 1986, Taiwan | Cross-sectional census survey | 12 | 0.6 ppm | 0.08 ppm | % with CDC | %, variance not required as census | No | Not applicable | Moderate | Yes |
| Ast and Chase, 1953, USA | Cross-sectional census survey | 6 | 1.2 ppm | 0.1 ppm | % with CDC | %, variance not required as census | No | Not applicable | Low | Yes |

Table 19 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF on the prevalence of mild to severe dental fluorosis

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|-----------------------------------|--|---|--------------------|---|--|--|--------------------------------------|---|---------------|----------------------------|
| Medcalf 1975, Australia | Cross-sectional survey | Samples of school children (6–8yrs) were examined pre and 6 years post fluoridation. None of the 1973 group has had lifetime exposure to fluoridation | 0.7–0.9 ppm | Pre-fluoridation in the goldfields (0.1–0.2 milligrams of fluoride per litre) | Dean's Index of fluorosis | No 95% CI for prevalence estimate | No | Not applicable | Low | Yes |
| Riordan and Banks 1991, Australia | Cross-sectional survey | School children aged born in 1978 Percentage lifetime exposure calculated | 0.8 ppm | Bunbury region fluoride <0.2 ppm | Thylstrup-Fejerskov index 0.78 | No 95% CI for prevalence estimate Yes 95% CI for ORs derived from regression analysis | Logistic regression analysis | Socio-economic status, areas of residence, type of regular water supply, and dates of changes since birth; periods and duration of use of fluoride supplements; use of fluoride toothpaste, including age commenced; and parent's recollection of whether the child liked and/or swallowed toothpaste | Low | Yes |
| Cortes <i>et al.</i> 1996, Brazil | Cross-sectional survey | School children 6–12-year-old lifetime residents using local drinking water sources from | 0.7 ppm | Maceio Alagoas <than 0.1 ppm of natural fluoride | Thylstrup-Fejerskov index | No prevalence calculation | No | Not applicable | Low | Yes |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|------------------------------------|---|--|--------------------|---|--|--|--------------------------------------|---|---------------|--|
| Heintze <i>et al.</i> 1998, Brazil | Cross-sectional survey | three economically deprived groups Participants aged 5–50 years were examined in health centres, schools, and factories | 0.75–1.2 ppm | Itapolis 0.02 ppm natural fluoridation | Thylstrup-Fejerskov index 0.85 | No 95% CI for prevalence estimate | No | Not applicable | Low | No, exclude on quality |
| Silva <i>et al</i> 2021, Brazil | Cross-sectional survey and adjusted for cluster sampling using a design effect of 1.7 | Children aged 5 years (daycare) and 12 years (school) | 0.5–0.6 ppm | Areas of Teresina not connected to piped water supply (< 0.05 ppm) | Thylstrup-Fejerskov index 0.90 | No 95% CI for prevalence estimate, but have design effect Yes 95% CI for ORs derived from regression analysis | Logistic regression analysis | Age, gender, lifetime exposure, socio-economic status, mother's level of education Snacks (sugar ingestion), CWF, fluoride toothpaste and toothbrushing Assess visits to dentist. | High | Consider for prevalence estimate Consider for independent contribution Divided into two groups very mild and mild and moderate so cannot use regression data |
| Brown 1951, Canada | Cross-sectional survey | School children at least 6 but not more than 14 years of age, not absent from the city concerned, for 'holidays or other reason, for more than six weeks at any one time | 1.0–1.2 ppm | Sarnia is fluorine-free, Stratford contains 1.3 ppm of fluorine from a natural source | Unidentified fluorosis index | No 95% CI for prevalence estimate | No | Not applicable | Low | No, exclude on quality |
| Brown 1960, Canada | Cross-sectional survey | Aged 9-11 and 12-14 years, 'continuous' residence, defined as | 1.0–1.2 ppm | Sarnia (fluorine-free, negligible amount of fluoride) and Stratford (1.3 ppm. | Unidentified fluorosis index | No prevalence calculation | No | Not applicable | Moderate | No, no prevalence calculation |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|---------------------------------------|--|---|--------------------|--|--|---|--------------------------------------|---|---------------|---|
| | | including absences, since birth, of six weeks or less. Residence eligibility is determined from information supplied by the parents. All schools of each city are canvassed | | of fluorine from a natural source) | | | | | | |
| <i>Brown and Poplove</i> 1965, Canada | Cross-sectional survey | All 16- and 17-year-old school children continuously resident in each city | 1.0–1.2 ppm | Sarnia (fluorine-free, negligible amount of fluoride) and Stratford (1.3 ppm. of fluorine from a natural source) | Unidentified fluorosis index | No prevalence calculation | No | Not applicable | Low | No, exclude on quality |
| Connor 1963, Canada | Cross-sectional census survey | Age groups 6, 7, 8; 9, 10, 11; and 12, 13, 14-year-old school children, continuous residents in each area | 1.0 ppm | Fluoride deficient survey in 1955 no baseline ppm reported but say water was fluoride free | Unidentified fluorosis index | Variance not required No prevalence calculation | No | Not applicable | Low | No, exclude on quality |
| Ismail <i>et al.</i> 1990, Canada | Cross-sectional survey and adjusted for cluster sampling but design effect unknown | Representative sample of public and private school students 11–17 years of age residing in Sherbrooke and Trois Rivières, Canada, who were born and lived at least the | 0.6 – 1.3 ppm | Sherbrooke, Quebec 0.1 ppm | Tooth Surface Index of Fluorosis 0.85 | Prevalence estimate with 95% CIs Yes 95% CI for ORs derived from regression analysis | Logistic regression analysis | Age, sex, residence, use of fluoridation supplement, type of school | Moderate | Consider for prevalence estimate Consider for independent contribution |

| Author, year, country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|---|--|--|--------------------|---|--|---|---------------------------------------|--|---------------|---|
| Clark <i>et al.</i> 1993, Canada | Cross-sectional survey | first 6 years of their life in their respective city Primary school-aged children, stratified by socio-economic status, residing in the respective cities, and had questionnaires completed | 1.2 ppm | Vernon fluoride deficient < 0.1 ppm | Tooth Surface Index of Fluorosis 0.44 | No 95% CI for prevalence estimate | No | Not applicable | Low | No, exclude on quality |
| <i>Clark et al.</i> 1994, Canada | Cross-sectional survey | All children aged 6–14 years [in selected schools] were asked to participate and randomly selected for inclusion, stratified by socio-economic status | 1.11 ppm | Vernon fluoride deficient <0.1 ppm | Tooth Surface Index of Fluorosis 0.44 | No 95% CI for ORs derived from regression analysis | Logistic regression | Fluoridated water, infant formula, use of fluoride supplementation and fluoride toothpaste during their first 6 years of life | Low | No, exclude on quality |
| <i>Ismail et al.</i> 1993, Canada | Cross-sectional census survey | School children in grades 5 and 6 in the two towns were included. Age not reported but children were over 6 years old maybe 10 to 12 years old). | 1.1 ppm | Truro, Nova Scotia, fluoride deficient <0.1 ppm | Tooth Surface Index of Fluorosis 0.90 | Variance not required for prevalence estimates No ORs for results of regression analysis | Stepwise multiple regression analysis | Sources of drinking water used since birth, residence history, gender, parents' education, use of fluoride supplements, toothpaste, and other fluoride products during their first 6 years of life | Moderate | No, exclude as numeric data were not provided for regression analysis |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|------------------------------------|--|--|-----------------------------------|---|--|--|---------------------------------------|---|---------------|---|
| Maupomé <i>et al.</i> 2003, Canada | Cross-sectional census survey | All of the school children examined who were lifelong residents in these communities | 0.88 ± 0.28 ppm - 0.92 ± 0.21 ppm | Comox/Courtenay and Campbell River stopped CWF in 1992 [fluoridation-ended (FE) communities] (FE 0.0 ppm) 14 to 19 months earlier | Thylstrup-Fejerskov index >0.75 | Variance not required for prevalence or regression as census survey | See Clark <i>et al.</i> 2006 | See Clark <i>et al.</i> 2006 | High | See Clark <i>et al.</i> 2006 |
| Clark <i>et al.</i> 2006, Canada | Cross-sectional census survey | Schoolchildren in 2nd and 3rd grades in 1993–94, 1996–97 and 2002–03, who were permanent residents | 0.88 ± 0.28 ppm - 0.92 ± 0.21 ppm | 2002–03 none of the children in had exposure to CWF (0.0 ppm) | Thylstrup-Fejerskov index 0.63 | Variance not required for prevalence or regression as census survey | Multivariate Poisson regression model | Residence history, parents' level of education, use of bottled water, consumption of breastmilk, infant formula, cow's milk and solid food, existence of home filtration devices, use and frequency of fluoride supplements, and/or mouth rinses. age brushing started, amount of toothpaste used, and brushing frequency | Moderate | No, exclude as numeric data were not provided for regression analysis |
| McLaren <i>et al.</i> 2021, Canada | Cross-sectional survey and adjusted for cluster sampling but design effect unknown | Grade 2 schoolchildren (~ age 7 years) enrolled in Public or Separate school systems in cities of Calgary and Edmonton | 0.59–0.89 ppm | Calgary CWF ceased in 2011 when levels of fluoride declined to 0.07–0.30 ppm | Tooth Surface Index of Fluorosis ≥ 0.80 | Prevalence estimate with 95% Cis Yes 95% CI for ORs derived from regression analysis | Poisson and logistic regression | Child's oral health, tooth brushing twice/day, floss once/day, visit dentist, last time visit dentist, dental insurance, fruits and vegetables at least once/day, Sugary drinks, fluoride | High | No, exclude as numeric data were not provided for regression analysis |

| Author, year, country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--|--|--|--------------------|---|--|--|--------------------------------------|--|---------------|--|
| | | | | | | | | supplements at home, fluoride treatments at the dentist's office, fluoride treatments in school programme, use of fluoride toothpaste, and/or fluoride mouth wash, household education, home ownership, and ethnocultural background | | |
| Villa <i>et al.</i> 1998a, Chile | Cross-sectional survey | 7-, 12-, and 15-year-old public or private schoolchildren who were lifelong residents of 5 areas | 0.93 ppm | Rancagua (0.7 ppm), Santiago (0.21 ppm natural), La Serena (0.55 ppm natural), and Iquique (1.10 ppm natural) | Dean's Index of fluorosis | No 95% CI for prevalence estimate | No | Age, social economic status | Moderate | No, as 95% CI data for prevalence estimate were not reported |
| Kunzel 1982, Cuba | Cross-sectional census survey | Children resident in study area | 0.7±0.1 ppm | The natural content of 0.05–0.1 ppm | Dean's Index of fluorosis | Variance not required for prevalence as census | No | Not applicable | Low | No, exclude on quality |
| Tabari <i>et al.</i> 2000, England, UK | Cross-sectional survey | 8–9-year-old school children who were lifetime residents in the area | 1.0 ppm | South Northumberland < 0.1 ppm | Thylstrup-Fejerskov index 0.70 | No 95% CI for prevalence estimate, but have design effect Yes 95% CI for ORs derived from regression analysis | Logistic regression analysis | Age started brushing, brushing frequency, type of toothpaste, amount of toothpaste, toothpaste weight, and socioeconomic class | Moderate | Consider for independent contribution |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--|---|---|--------------------------|---|--|---|--|---|---------------|--|
| Ellwood and O'Mullane 1996, England, Wales, UK | Cross-sectional survey Census for intervention group | School children in the third year of their secondary school education, who were lifetime residents of the areas | 0.7 ppm | Chester (England) and Bala (North Wales) < 0.1 ppm | Thylstrup-Fejerskov index 0.73 | No 95% CI for comparator prevalence estimate | No | Not applicable | Low | No, based on quality and 95% CI data for prevalence estimate were not reported |
| Clarkson and O'Mullane 1992, Ireland | Cross-sectional survey | 8-year-old school children | 0.8–1.0 ppm | Fluoride deficient water in Ireland is ≤0.3 ppm | Dean's Index of fluorosis | No 95% CI for prevalence estimates | No | Not applicable | Low | No, based on quality and 95% CI data for prevalence estimate were not reported |
| Whelton <i>et al.</i> 2004, Ireland | Cross-sectional survey and description indicate that authors have adjusted for cluster sampling but not stated it | 5-, 8-, 12- and 15-year-old school children living in the Republic of Ireland | 0.8–1.0 ppm | Not reported (fluoride deficient parts of Ireland are ≤0.3 ppm) | Dean's Index of fluorosis | No 95% CI for prevalence estimates | No | Not applicable | Moderate | No, as 95% CI data for prevalence estimate were not reported |
| James <i>et al.</i> 2021, Ireland | Cross-sectional survey and description indicate that authors have adjusted for cluster sampling but not stated it | Random sample of 5-year-old schoolchildren in Dublin & Cork-Kerry in 2014, follow up at age 8 years in 2017 | 0.8–1.0 ppm | Fluoride deficient County Cork and Kerry ≤0.3 ppm | Dean's Index of fluorosis 0.74 | No 95% CI for prevalence estimates Yes 95% CI for ORs derived from regression analysis | Multivariate regression with a negative binomial Hurdle model for caries only Logistic regression for fluorosis. Both comparing the difference in two time points | Age, gender, age first used toothpaste, amount of toothpaste, frequency of toothbrushing, age first visited dentist, rinse method after toothbrushing, and sweet snacks between meals | High | No, as 95% CI data for prevalence estimate were not reported and do not provide individual contributions and are tests of difference |
| Mohd Nor <i>et al.</i> 2018, Malaysia | Cross-sectional survey and description indicate that | School children aged 9 (born 2006) and 12 (born 2003), | 0.7 ppm and then 0.5 ppm | Kelantan described as fluoride deficient (0 ppm) confirmed | Dean's Index of fluorosis 0.72–0.90 | Prevalence estimate with 95% CIs | See Mohd Nor <i>et al.</i> 2021 | See Mohd Nor <i>et al.</i> 2021 | Moderate | No as not sure if adjusted for cluster |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|---------------------------------------|--|---|--------------------------|--|--|---|--------------------------------------|--|---------------|---------------------------------------|
| | authors have adjusted for cluster sampling but not stated it | lifelong residents were included in the final analysis | | | | Query on adjustment for cluster sampling | | | | sampling for 95% CIs |
| <i>Mohd Nor et al.</i> 2021, Malaysia | Cross-sectional survey (as above) | School children aged 9 (born 2006) and 12 (born 2003), lifelong residents were estimated for this study | 0.7 ppm and then 0.5 ppm | Kelantan described as fluoride deficient (0 ppm) confirmed | Dean's Index of fluorosis 0.72–0.90 | Yes 95% CI for ORs derived from regression analysis | Multiple binary logistic regression | CWF, filter, and bottled water use, infant feeding patterns (breast and formula feeding), oral hygiene practices at age less than 6 years (age at which the child started toothbrushing with toothpaste, toothbrushing supervision, frequency of brushing, behaviour after brushing, habits of eating and licking toothpaste, amount and the type of toothpaste used), socioeconomic, and demographic background | Moderate | Consider for independent contribution |
| Ministry of Health 2010, New Zealand | Cross-sectional survey and adjusted for cluster sampling using a design effect of ≥ 2 | In households, one adult aged ≥ 15 years, and one child aged from birth to 14 years old, if any, were randomly selected for the survey | 0.8–0.9 ppm | Circa 0.15 ppm in fluoride deficient areas | Dean's Index of fluorosis 0.78 | Prevalence estimate with 95% CIs Query control for lifetime exposure as estimate higher in | No | Not applicable | Moderate | Consider for prevalence estimate |

| Author, year, country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|------------------------------------|--|---|--------------------------|---|--|--|--------------------------------------|--|---------------|---|
| | | | | | | fluoride deficient area | | | | |
| Wong <i>et al.</i> 1970, Singapore | Cross-sectional survey | Chinese and Malay children in two age groups, 7–8 years and 8–9 years of age were selected by random sampling from primary schools in various parts of the island | 0.7 ppm | Before fluoridation was 0.2 ppm | No index used. Dental fluorosis observed | No 95% CI for prevalence estimates | No | Not applicable | Low | No, based on quality and 95% CI data for prevalence estimate were not reported |
| Hong <i>et al.</i> 1990, Taiwan | Cross-sectional census survey | Children aged 6–15 years who were born in or continuous residents of the respective areas | 0.6 ppm and then 0.7 ppm | Tsao-tun (0.08 ppm) | Dean's Index of fluorosis | Variance not required for prevalence as census | No | Not applicable | Moderate | Consider for prevalence estimate |
| Arnold <i>et al.</i> 1956, USA | Cross-sectional survey | Kindergarten and school children aged 4-16 years who had used city water supplies continuously since birth | 0.9–1.1 ppm | Muskegon, <0.2 ppm of fluoride until July 1951, 1952 - 1954 1.0 ppm, Aurora Natural F 1.2 ppm | No index used. Proportion with dental fluorosis observed | No 95% CI for prevalence estimates | No | Not applicable | Low | No, based on quality and 95% CI data for prevalence estimate were not reported |
| Szpunar and Burt 1988, USA | Cross-sectional survey | 6–12-year-old school children | 1.0 ppm | Natural fluoride: Richmond 1.2 ppm, Cadillac 0.0 ppm, Hudson 0.8 ppm, fluoride mouth rinses | Tooth Surface Index of Fluorosis 0.85 | No 95% CI for prevalence estimates Yes, 95% CI for ORs derived from regression analysis | Logistic regression analysis | Demographic information, residence history, details of fluoride exposure, information about the use of dental services, and infant nutrition | Low | No, based on quality and as 95% CI data for prevalence estimate were not reported |

| Author, year country | Study design and census/cluster sample adjustment where reported | Study population and lifetime exposure | CWF exposure (ppm) | Details of comparator | Fluorosis outcome measure and proportion agreement where reported cut offs | Statistical measure and variance | Regression to adjust for confounding | Confounders | Study quality | Consider for meta-analysis |
|--------------------------------------|--|---|--------------------|---|--|--|--------------------------------------|--|---------------|--|
| Kumar <i>et al.</i> 1989, USA | Cross-sectional survey and adjusted for cluster sampling but design effect unknown | 7–14-year-old school children. Children with orthodontic bands, only deciduous teeth, not lifetime residents of respective cities were excluded | 1.0 ppm | Kingston < 0.3 ppm | Dean's Index of fluorosis | No 95% CI for prevalence estimates | No | Not applicable | Low | No, based on quality and 95% CI data for prevalence estimate were not reported |
| <i>Kumar et al.</i> 1998, USA | Cross-sectional survey and adjusted for cluster sampling but design effect unknown | School children in grades 1 through 8, children aged 7 to 14 years, who had been lifelong residents of respective cities. | 1 ± 0.2 ppm | Kingston < 0.3 ppm | Dean's Index of fluorosis 0.65, 0.76, and 1.0 for 3 of the examiners relative to the fourth | No 95% CI for prevalence estimates | No | Not applicable | Low | No, based on quality and 95% CI data for prevalence estimate were not reported |
| <i>Kumar et al.</i> 2000, USA | Cross-sectional survey and adjusted for cluster sampling but design effect unknown | School children who were 7–14-year-old lifelong residents | 1 ± 0.2 ppm | Kingston, New Windsor, Town of Ulster < 0.3 ppm | Dean's Index of fluorosis 0.65, 0.76, and 1.0 for 3 of the examiners relative to the fourth | Yes, 95% CI for ORs derived from regression analysis | Logistic regression | Age, sex, ethnic group, socioeconomic status, use of fluoride tablets/drops during the first eight years of life | Low | No, based on quality |

6.10 Appendix J Tables used for the meta-analyses presented in the main report

Table 20 Primary dentition: dental caries measured using dmft in CWF areas compared with fluoride-deficient areas (sensitivity analysis with three outlier papers removed)

| Author | Year | Country | Age in years | CWF ppm | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF Mean | CWF SD | Fluoride deficient participants | Fluoride deficient Mean | Fluoride deficient SD |
|-------------------------|-------|-------------|--------------|-----------|-----------------------|------------------------|----------|---------------------|------------------|----------|-----------|---------------------------------|-------------------------|-----------------------|
| French et al. | 1984 | England | 5 | 1.0 | 3 | 0.1 | Low | Backer-Dirks et al. | 533 | 1.41 | 2.21 | 536 | 3.37 | 3.65 |
| Jackson et al. | 1975b | England | 5 | 1.0 | 3 | <0.1 | Low | Jackson et al. | 106 | 2.38 | 0.304 SE± | 130 | 4.4 | 0.349 SE± |
| Lemasney et al. | 1984 | Ireland | 5 | 0.8 – 1.0 | 3 | ≤0.3 | Low | Whittle and Downer | 169 | 2.46 | 3.27 | 98 | 3.83 | 3.75 |
| Rugg-Gunn et al. | 1977 | England | 5 | 1.0 | 3 | <0.1 | Low | Backer-Dirks et al. | 212 | 2.4 | 2.73 | 132 | 6.1 | 4.03 |
| Rugg-Gunn et al. | 1981 | England | 5 | 1.0 | 3 | <0.1 | Low | Backer-Dirks et al. | 438 | 2.5 | 2.79 | 132 | 6.1 | 4.03 |
| Seaman et al. | 1989 | Wales | 5 | 0.8 | 2 | <0.1 | Low | Palmer et al. | 260 | 0.8 | 1.43 | 546 | 2.26 | 3.17 |
| Guo et al. 1984 | 1984 | Taiwan | 5 | 0.6 | 2 | 0.08 | Moderate | WHO | 345 | 5.5 | 4.3 | 387 | 8.5 | 4.6 |
| Hsieh et al. 1986 | 1986 | Taiwan | 5 | 0.6 | 2 | 0.08 | Moderate | WHO | 226 | 5.1 | 3.8 | 319 | 8.6 | 4 |
| O'Mullane et al. 1986 | 1986 | England | 5 | 0.8–1.0 | 3 | ≤3 | Moderate | WHO | 869 | 1.8 | 2.8 | 836 | 3 | 3.7 |
| Rugg-Gunn et al. 1988 | 1988 | England | 5 | 1 | 3 | <0.1 | Moderate | Backer-Dirks | 457 | 1.81 | 2.56 | 370 | 3.9 | 4.22 |
| Treasure and Dever 1992 | 1992 | New Zealand | 5 | 1 | 3 | 0.08 | Moderate | Palmer | 107 | 1.06 | 1.75 | 67 | 2.91 | 3.82 |
| Evans et al. 1995 | 1995 | England | 5 | 1 | 3 | <0.1 | Moderate | BASCD | 496 | 1.33 | 0.57 | 436 | 2.41 | 0.53 |
| Villa et al. 1998 | 1998 | Chile | 7 | 0.93 | 3 | 0.07 | Moderate | WHO | 129 | 1.72 | 2.33 | 158 | 3.67 | 3.54 |

| | | | | | | | | | | | | | | |
|---------------------|------|---------|---|-----------------|---|-----------|----------|-----------------------|------|------|------|------|------|------|
| Whelton et al. 2004 | 2004 | Ireland | 5 | 0.8–1.0 | 3 | ≤0.3 | Moderate | WHO | 3616 | 1 | 2.1 | 2160 | 1.7 | 2.1 |
| James et al. 2021 | 2021 | Ireland | 8 | 0.8–1.0–0.6–0.8 | 3 | ≤0.3 | High | WHO | 704 | 1.9 | 2.4 | 770 | 2.7 | 2.8 |
| McLaren et al. 2021 | 2021 | Canada | 7 | 0.6–0.8 | 2 | 0.07–0.30 | High | WHO | 799 | 2 | 4.33 | 918 | 3.2 | 3.86 |
| Silva et al. 2021 | 2021 | Brazil | 5 | 0.5–0.6 | 2 | <0.05 | High | Not reported | 161 | 1.53 | 2.47 | 169 | 3.54 | 4.1 |
| Goodwin et al. 2022 | 2022 | England | 5 | 0.9 | 3 | <0.2 | Moderate | Public Health England | 699 | 1.06 | 2.16 | 911 | 1.18 | 2.41 |

Table 21 Primary dentition: dental caries measured using dmfs in CWF areas compared with fluoride-deficient areas (sensitivity analysis with one outlier paper removed)

| Author | Year | Country | Age in years | CWF ppm | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF Mean | CWF SD | Fluoride deficient participants | Fluoride deficient Mean | Fluoride deficient SD |
|-------------------------|-------|-------------|--------------|---------|-----------------------|------------------------|----------|---------------------|------------------|----------|--------|---------------------------------|-------------------------|-----------------------|
| Rugg-Gunn et al. | 1977 | England | 5 | 1 | 3 | <0.1 | Low | Backer-Dirks et al. | 212 | 3.6 | 4.98 | 132 | 11.6 | 9.54 |
| Rugg-Gunn et al. | 1981 | England | 5 | 1 | 3 | <0.1 | Low | Backer-Dirks et al. | 438 | 4.1 | 5.76 | 132 | 11.6 | 9.54 |
| French et al. | 1984 | England | 5 | 1 | 3 | <0.1 | Low | Backer-Dirks et al. | 533 | 2.14 | 4.13 | 536 | 5.7 | 7.19 |
| Seppa et al. | 2000b | Finland | 6 | 1 | 3 | <0.1 | Low | Moller & Poulsen | 49 | 2.53 | 3.1 | 66 | 1.32 | 2.51 |
| Rugg-Gunn et al. 1988 | 1988 | England | 5 | 1 | 3 | <0.1 | Moderate | Backer-Dirks | 457 | 2.8 | 4.77 | 370 | 7 | 9.28 |
| Treasure and Dever 1992 | 1992 | New Zealand | 5 | 1 | 3 | NR | Moderate | Palmer | 107 | 1.52 | 2.65 | 67 | 4.69 | 7.03 |

Table 22 Primary dentition: Percent with non cavitated dental caries measured using dmft in CWF areas compared with fluoride-deficient areas

| Author | Year | Country | Age in years | CWF population | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF % with CD | CWF 95% CI | Fluoride deficient participants | Fluoride deficient % with CDC | Fluoride deficient 95% CI | % Difference |
|------------------------|------|---------|--------------|----------------|-----------------------|------------------------|-----------|--|------------------|---------------|-------------|---------------------------------|-------------------------------|---------------------------|--------------|
| Brown et al. | 1960 | Canada | 9-11 | 1.0-1.2 | 3 | NF | Mode rate | Not reported | 502 | 41.83 | 2.20 2SE | 521 | 34.36 | 2.081 SE | 7.47 |
| Gray and Davies-Slowik | 2001 | England | 5 | 1 | 3 | <0.3 | Low | British Association for the Study of Community Dentistry | 379 | 79.8 | 79.4 – 82.0 | 273 | 65.2 | 64.6–67.5 | 14.6 |
| Gray and Davies-Slowik | 2001 | England | | | | | | British Association for the Study of Community Dentistry | 413 | 69.5 | 69.1 – 71.7 | 273 | 65.2 | 64.6–67.5 | 4.3 |
| Gray and Davies-Slowik | 2001 | England | | | | | | British Association for the Study of Community Dentistry | 660 | 74.1 | 73.8 – 76.2 | 273 | 65.2 | 64.6–67.5 | 8.9 |
| Gray and Davies-Slowik | 2001 | England | | | | | | British Association for the Study of Community Dentistry | 451 | 80 | 79.6 – 82.2 | 273 | 65.2 | 64.6–67.5 | 14.8 |
| Gillcrist et al. | 2001 | USA | 5-11 | 1 | 3 | <0.3 | Low | ADA | 10,495 | 42 | 39–44 | 6,761 | 35 | 32–37 | 7 |

| Author | Year | Country | Age in years | CWF ppm | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF % with CDC | CWF 95% CI | Fluoride deficient participants | Fluoride deficient % with CDC | Fluoride deficient 95% CI | % Difference |
|---------------|------|---------|--------------|---------|-----------------------|------------------------|---------|-------|------------------|----------------|------------|---------------------------------|-------------------------------|---------------------------|--------------|
| Ast and Chase | 1953 | USA | 5 | 1.2 | 3 | <0.1 | Low | WHO | 217 | 56.2 | 0.01 | 140 | 26.4 | 0.01 | 29.8 |

Table 23 Primary dentition: Percent with cavitated dental caries measured using dmft in CWF areas compared with fluoride-deficient areas

| Author | Year | Country | Age in years | CWF ppm | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF participants with CDC | CWF % with CDC | CWF low CI | CWF upper CI | Fluoride deficient participants | Fluoride deficient participants with CDC | Fluoride deficient % with CDC | Fluoride deficient lower CI | Fluoride deficient upper CI | % Difference |
|---------------------|------|---------|--------------|---------|-----------------------|------------------------|----------|-------|------------------|---------------------------|----------------|------------|--------------|---------------------------------|--|-------------------------------|-----------------------------|-----------------------------|--------------|
| Guo et al. 1984 | 1984 | Taiwan | 5 | 0.6 | 2 | 0.08 | Moderate | WHO | 345 | 298 | 86.4 | 0.1 | 0.1 | 387 | 368 | 95.1 | 0.1 | 0.1 | 8.7 |
| Hsieh et al. 1986 | 1986 | Taiwan | 5 | 0.6-0.7 | 2 | 0.08 | Moderate | WHO | 226 | 225 | 99.6 | 0.1 | 0.1 | 319 | 318 | 99.7 | 0.1 | 0.1 | 0.1 |
| Evens et al. 1995 | 1995 | England | 5 | 1 | 3 | <0.1 | Moderate | BASCD | 496 | 193 | 39 | 0.1 | 0.1 | 436 | 240 | 55 | 0.1 | 0.1 | 16 |
| McLaren et al. 2021 | 2021 | Canada | 7 | 0.6-0.8 | 2 | 0.07-0.30 ppm | High | WHO | 799 | 356 | 44.5 | 44.5 | 49.2 | 918 | 558 | 60.8 | 57 | 64.5 | 16.3 |

Table 24 Permanent dentition DMFT in CWF areas compared with fluoride-deficient areas (sensitivity analysis with 4 outlier papers removed)

| Author | Year | Country | Age in years | CWF ppm | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF Mean | CWF SD | Fluoride deficient participants | Fluoride deficient Mean | Fluoride deficient SD |
|------------------------|------|-------------|--------------|-----------|-----------------------|------------------------|----------|-----------------------|------------------|----------|--------|---------------------------------|-------------------------|-----------------------|
| de Liefde and Herbison | 1985 | New Zealand | 9 | 1.0 | 3 | 0. | Low | WHO | 191 | 1.7 | 1.6 | 237 | 2.4 | 1.9 |
| Kunzel | 1980 | Germany | 10 | 1.0 | 3 | 0.07 | Low | Not reported | 164 | 1.3 | 1.41 | 272 | 3.1 | 1.95 |
| Lemasney et al. | 1984 | Ireland | 11 | 0.8 – 1.0 | 3 | <0.1 | Low | Whittle and Downer | 182 | 2.12 | 1.97 | 126 | 3.63 | 2.79 |
| Kunzel et al., | 2000 | Germany | 12 | 0.8 – 1.0 | 3 | 0.05-0.1 | Low | WHO | 337 | 2.47 | 2.06 | 472 | 4.65 | 1.77 |
| Mitropoulos et al. | 1988 | England | 14 | 1.0 | 3 | NF | Low | Downer et al. | 234 | 2.26 | 2.46 | 275 | 3.79 | 3.22 |
| Kalsbeek et al., | 1993 | Netherlands | 15 | 1.1 | 3 | <0.1 | Low | Modified Backer Dirks | 285 | 7.4 | 4 | 261 | 14.1 | 5.7 |
| Clovis et al. | 1988 | Canada | 11-12 | 1.08 | 3 | 0.23 | Low | WHO | 53 | 2.26 | 2.43 | 77 | 2.43 | 2.11 |
| Murray et al. | 1991 | England | 15-16 | 1.0 | 3 | 0.07–0.30 | Low | Palmer et al. | 349 | 2.7 | 0.13 | 347 | 3.4 | 0.16 |
| Hsieh et al. 1979 | 1979 | Taiwan | 6 | 0.6 – 0.7 | 2 | 0.08 | Moderate | WHO | 312 | 0.1 | 0.4 | 238 | 0.3 | 0.7 |
| Guo et al. 1984 | 1984 | Taiwan | 10 | 0.6 | 2 | 0.08 | Moderate | WHO | 310 | 1.1 | 1.5 | 436 | 2.4 | 2 |
| Hsieh et al. 1986 | 1986 | Taiwan | 12 | 0.6 – 0.7 | 2 | <0.1 | Moderate | WHO | 329 | 1.9 | 2.4 | 458 | 4.3 | 3.6 |
| O'Mullane et al. 1986 | 1986 | Ireland | 12 | 0.8 – 1.0 | 3 | 0 | Moderate | Index | 749 | 2.6 | 2.3 | 755 | 3.3 | 2.5 |
| Thomas and Kassab 1992 | 1992 | Wales | 18-30 | 0.8 | 2 | <0.1 | Moderate | WHO | 170 | 9.48 | 4.04 | 479 | 13.62 | 4.6 |

| Author | Year | Country | Age in years | CWF ppm | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF Mean | CWF SD | Fluoride deficient participants | Fluoride deficient Mean | Fluoride deficient SD |
|-------------------------|------|-------------|--------------|-----------|-----------------------|------------------------|----------|-------|------------------|----------|--------|---------------------------------|-------------------------|-----------------------|
| Treasure and Dever 1994 | 1994 | New Zealand | 14 | 1 | 3 | 0.08 | Moderate | WHO | 134 | 2.33 | 2.16 | 48 | 4.52 | 3.7 |
| Villa et al. 1998 | 1998 | Chile | 12 | 0.93 | 3 | ≤3 | Moderate | WHO | 152 | 1.28 | 1.65 | 155 | 3.1 | 2.65 |
| Kunzel and Fischer 2000 | 2000 | Cuba | 10-11 | 0.8 | 2 | <0.3 | Moderate | Index | 126 | 1.1 | 1.51 | 85 | 3.1 | 1.79 |
| Whelton et al. 2004 | 2004 | Ireland | 12 | 0.8 – 1.0 | 3 | 0.1 | Moderate | WHO | 2090 | 1.1 | 1.4 | 747 | 1.3 | 1.7 |
| Mullen et al. 2012 | 2012 | Ireland | 16 | 0.7 | 2 | 0.2 | Moderate | WHO | 823 | 2.42 | 4.46 | 253 | 3.61 | 2.03 |
| Mohd Nor et al. 2018 | 2018 | Malaysia | 12 | 0.5 | 1 | 0 | Moderate | WHO | 294 | 0.47 | 0.97 | 301 | 1.31 | 1.81 |
| McLaren et al. 2021 | 2021 | Canada | 7 | 0.6 – 0.8 | 2 | ≤0.3 | High | Index | 791 | 0.19 | 0.78 | 912 | 0.26 | 1 |
| Silva et al. 2021 | 2021 | Brazil | 12 | 0.6 | 2 | <0.05 | High | WHO | 178 | 1.53 | 1.81 | 184 | 2.63 | 3.02 |

Table 25 Permanent dentition DMFT in CWF areas compared with fluoride-deficient areas (sensitivity analysis with 1 outlier paper removed)

| Author | Year | Country | Age in years | CWF ppm | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF Mean | CWF SD | Fluoride deficient participants | Fluoride deficient Mean | Fluoride deficient SD |
|-----------------------|------|---------------|--------------|---------|-----------------------|------------------------|---------|-----------------------|------------------|----------|---------------------|---------------------------------|-------------------------|-----------------------|
| Ellwood and O'Mullane | 1995 | England Wales | 14 | 0.7 | 2 | <0.1 | Low | Stephen et al. | 196 | 3.18 | 3.92 | 267 | 4.18 | 4.56 |
| Kalsbeek et al. | 1993 | Netherlands | 15 | 1.1 | 3 | 0.1 | Low | Modified Backer Dirks | 285 | 10.8 | 7.7 | 261 | 27.7 | 14.6 |
| Gillcrist et al. | 2001 | USA | 5-11 | 1.0 | 3 | <0.3 | Low | ADA | 10,495 | 0.77 | 0.65, 0.88 (95% CI) | 6,761 | 1.02 | 0.90, 1.13 (95% CI) |

| Author | Year | Country | Age in years | CWF ppm | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF Mean | CWF SD | Fluoride deficient participants | Fluoride deficient Mean | Fluoride deficient SD |
|--------------------|------|-------------|--------------|---------|-----------------------|------------------------|----------|-------------|------------------|----------|--------|---------------------------------|-------------------------|-----------------------|
| Kunzel and Fischer | 2000 | Cuba | 10-11 | 0.8 | 2 | 0.05-0.1 | Moderate | WHO | 126 | 1.5 | 2.21 | 85 | 4.8 | 3.76 |
| Treasure and Dever | 1994 | New Zealand | 14 | 1.0 | 3 | 0.08 | Moderate | Palm et al. | 134 | 2.97 | 3.08 | 48 | 6.19 | 6.41 |

Table 26 Permanent dentition: Percent with non cavitated dental caries measured using dmft in CWF areas compared with fluoride-deficient areas

| Author | Year | Country | Age in years | CWF ppm | CWF exposure category | Fluoride deficient ppm | Quality | Index | CWF participants | CWF % with CDC | CWF 95% CI | Fluoride deficient participants | Fluoride deficient % with CDC | Fluoride deficient 95% CI | % Difference |
|--------------------|------|---------|--------------|---------|-----------------------|------------------------|----------|-------|------------------|----------------|------------|---------------------------------|-------------------------------|---------------------------|--------------|
| Brown et al. | 1960 | Canada | 12-14 | 1.0-1.2 | 3 | NF | Moderate | NR | 503 | 18.69 | 1.738 | 485 | 2.27 | 0.676 | 16.42 |
| Brown and Poplove, | 1965 | Canada | 16-17 | 1.0-1.2 | 3 | NF | Low | NR | 356 | 11.8 | 1.71 | 482 | 0.41 | 0.291 | 11.39 |
| Gillcrist et al., | 2001 | USA | 5-11 | 1.0 | 3 | <0.3 | Low | ADA | 10,495 | 78 | 76,80 | 6,761 | 74 | 72,76 | 4.0 |

Table 27 Permanent dentition: percent with cavitated dental caries measured using DMFT in CWF areas compared with fluoride-deficient areas

| Author | Year | Country | Age | CWF | CWF exposure category | Fluoride deficient population | Quarantine | Individual | CWF participants | CWF participants with CDC | CWF % with CDC | CWF % with CI | CWF % with CI | Fluoride deficient participants | Fluoride deficient participants with CDC | Fluoride deficient % with CDC | Fluoride deficient lower CI | Fluoride deficient upper CI | % Difference |
|---------------------|------|---------|-----|-----------|-----------------------|-------------------------------|------------|------------|------------------|---------------------------|----------------|---------------|---------------|---------------------------------|--|-------------------------------|-----------------------------|-----------------------------|--------------|
| Guo et al. 1984 | 1984 | Taiwan | 10 | 0.6 | 2 | 0.08 | Moderate | WHO | 310 | 149 | 48.1 | 0.01 | 0.01 | 436 | 352 | 80.7 | 0.01 | 0.01 | 0.326 |
| Hsieh et al. 1986 | 1986 | Taiwan | 12 | 0.6 | 2 | 0.08 | Moderate | WHO | 329 | 197 | 59.9 | 0.01 | 0.01 | 458 | 381 | 83.2 | 0.01 | 0.01 | 0.232 |
| McLaren et al. 2021 | 2021 | Canada | 7 | 0.6 - 0.8 | 2 | 0.07 - 0.30 | High | WHO | 791 | 98 | 12.4 | 9.6 | 15.9 | 912 | 141 | 15.4 | 12.4 | 18.9 | 0.03 |

7 Appendices Question 2A

7.1 Appendix A Overview of literature search results for Question 2 A and B

Table 28 Overview of literature search for Question 2A and 2B

| Database | Date of search | Date range | No. of results |
|--|----------------|----------------|----------------|
| Ovid MEDLINE(R) and Epub Ahead of Print, In-Process, In-Data-Review & Other Non-Indexed Citations, Daily and Versions | 07 Dec 2021 | 1946-2021 | 1866 |
| Embase | 07 Dec 2021 | 1974-2021 | 481 |
| Cochrane Library (John Wiley & Sons Inc) | 07 Dec 2021 | 1946-2021 | 51 |
| Cochrane Trial Register | 07 Dec 2021 | 1946-2021 | |
| LILACS | 27 July 2021 | Inception-2021 | 96 |
| Scoping search | | | 70 |
| Ovid MEDLINE(R) and Epub Ahead of Print, In-Process, In-Data-Review & Other Non-Indexed Citations, Daily and Versions (1946-1990) | 28 Feb 2023 | 2021-2023 | 93 |
| Embase | 28 Feb 2023 | 2021-2023 | 39 |
| Total before deduplication | | | 2696 |
| Total after deduplication | | | 2,223 |
| Total retained for analysis after screening Q2A | | | 16 |
| Total retained for analysis after screening Q2B | | | 4 |
| Total added from reference chasing Q2A | | | 3 |
| Total added from reference chasing Q2B | | | 3 |

7.1.1 Medline (1946-06 December 2021)

Database(s): Ovid MEDLINE(R) and Epub Ahead of Print, In-Process, In-Data-Review & Other Non-Indexed Citations, Daily and Versions(R) (1946 to December 06, 2021)

| # | Searches | Results |
|---|--|---------|
| 1 | exp Fluoride/ | 38309 |
| 2 | (fluorid* or fluorin* or flurid* or florin*).ti. or (fluorid* or fluorin* or flurid* or florin*).ab. or (fluorid* or fluorin* or flurid* or florin*).sh,kf,kw. | 97770 |
| 3 | 1 or 2 | 102195 |
| 4 | Water/ or water.mp. | 1052918 |

| | | |
|----|---|---------|
| 5 | Water Supply/ | 33625 |
| 6 | 4 or 5 | 1052918 |
| 7 | 3 and 6 | 15053 |
| | exp adolescent/ or exp child/ or exp infant/ or (infant disease* or childhood disease*).ti,ab,kf. or (adolescen* or babies or baby or boy? or boyfriend or boyhood or child* or girl? or infant* or | |
| 8 | juvenil* or kid? or minors or minors* or neonat* or neonat* or newborn* or new-born* or paediatric* or peadiatric* or pediatric* or perinat* or preschool* or puber* or pubescen* or school* or teen* or toddler? or underage? or under-age? or youth*).ti,ab,kf. | 4712855 |
| 9 | exp Child Health/ or Child Health Services/ or exp Pediatrics/ | 84603 |
| 10 | 8 or 9 | 4718792 |
| 11 | exp Dental Caries/ | 48277 |
| 12 | carie\$.mp. | 63352 |
| 13 | (caries or early childhood caries).mp. [mp=title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms] | 63235 |
| 14 | exp Periodontal Diseases/ | 91326 |
| 15 | exp tooth demineralization/ | 49853 |
| 16 | (teeth adj5 (cavit\$ or caries\$ or carious or decay\$ or lesion\$ or deminerali\$ or reminerali\$)).mp. | 12198 |
| 17 | (tooth adj5 (cavit\$ or caries\$ or carious or decay\$ or lesion\$ or deminerali\$ or reminerali\$)).mp. | 10220 |
| 18 | (dental adj5 (cavit\$ or caries\$ or carious or decay\$ or lesion\$ or deminerali\$ or reminerali\$)).mp. | 73817 |
| 19 | (enamel adj5 (cavit\$ or caries\$ or carious or decay\$ or lesion\$ or deminerali\$ or reminerali\$)).mp. | 5802 |
| 20 | (dentin\$ adj5 (cavit\$ or caries\$ or carious or decay\$ or lesion\$ or deminerali\$ or reminerali\$)).mp. | 7488 |
| 21 | Dental Enamel/ | 20238 |
| 22 | (root\$ adj5 (cavit\$ or caries\$ or carious or decay\$ or lesion\$ or deminerali\$ or reminerali\$)).mp. | 6089 |
| 23 | ((teeth or tooth or dental or enamel or dentin) and plaque).mp. | 27678 |
| 24 | Oral Health/ | 18506 |
| 25 | (Life Quality or Health-Related Quality Of Life or Health Related Quality Of Life).mp. or HRQOL.ab,ti,kw. | 60264 |
| 26 | ("DMF Index" or "Dental Plaque Index").mp. | 14771 |
| 27 | or/11-26 | 266603 |
| 28 | exp Fluorides, Topical/ | 4744 |

| | | |
|----|--|---------|
| 29 | exp Fluorides/ | 38309 |
| 30 | Fluor\$.mp. | 1164236 |
| 31 | monofluor\$.mp. | 1599 |
| 32 | exp Cariostatic Agents/ | 36787 |
| 33 | (fluoride varnish or bifluorid or cavityshield or duraflur or duraphat or fluorniz or fluor protector or prevent varnish or thera-flur or clinpro white varnish).mp. [mp=title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms] | 1299 |
| 34 | (varnish* or lacquer* or laquer* or paint*).mp. [mp=title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms] | 29115 |
| 35 | exp Dentifrices/ | 7096 |
| 36 | (Mouth Bath or Mouth Wash or Mouth Rinse).ab,ti,kf. | 970 |
| 37 | (toothpaste\$ or paste\$ or dentrifice\$).mp. | 42569 |
| 38 | (varnish adj5 tooth).mp. | 79 |
| 39 | 28 or 29 or 30 or 31 or 32 or 33 or 34 or 35 or 36 or 37 or 38 | 1230993 |
| 40 | 7 and 10 and 27 and 39 | 1911 |
| 41 | exp animals/ not humans.sh. | 4924363 |
| 42 | 40 not 41 | 1901 |
| 43 | limit 42 to (comment or editorial or letter or newspaper article) | 36 |
| 44 | 42 not 43 | 1866 |

7.1.2 Embase (1974-06 December 2021)

| # | Searches | Results |
|---|--|---------|
| 1 | Fluorides/ or Fluorine/ | 38765 |
| 2 | exp Fluoridation/ | 4411 |
| 3 | water.mp. /freq=5 and (fluorid\$ or fluorin\$ or flourid\$ or flourin\$ or flurid\$ or flurin\$ or florid\$ or florin\$).mp. | 4890 |
| 4 | (Hexafluorsilicic acid or Hydrofluosilicic acid or HFSA or "H ₂ SiF ₆ " or "CaF ₂ " or fluorospar or fluorosilicic acid or sodium fluorosilicate\$ or silicofluorid\$).mp. | 1267 |
| 5 | or/1-4 | 44530 |
| 6 | Water Supply/ or Water/ or (drinking water or drinking suppl\$ or potable water or water suppl\$ or suppl\$ of water or public water or community water or water treatment or waterworks or water fluorid\$).mp. | 437124 |
| 7 | 5 and 6 | 8511 |
| 8 | (topical\$ adj5 fluor\$).mp. | 3365 |

| # | Searches | Results |
|----|---|---------|
| 9 | (Fluorides/ or Fluorine/) and Cariostatic agents/ | 2500 |
| 10 | exp Dentifrices/ or Mouthwashes/ or Toothpastes/ | 12520 |
| 11 | (toothpaste\$ or paste\$ or dentrifice\$).mp. | 53731 |
| 12 | fluoride prophylaxis toothpaste/ or fluoride varnish/ | 3457 |
| 13 | ((varnish\$ or gel or gels or rinsing or rinse) adj5 (dental or tooth or fluorid\$)).mp. | 5554 |
| 14 | (Mouth bath or mouthbath or mouth wash\$ or mouthwash\$ or Mouth Rinse\$ or mouthrinse\$).mp. | 8557 |
| 15 | ((Fluor\$ or AMF or APF or "Amine F" or SNF2 or "Stannous F" or NAF or "Sodium F" or SMFP or MFP or monofluor\$ or "PPM F" or PPMF or "phosphat\$ F" or "acidulat\$ Fluor\$" or "phosphat\$ fluor\$" or fluorphosphat\$ or "amin\$ fluor\$" or "sodium fluor\$") and (topical\$ or paste\$ or gel or gels or varnish\$ or administration route\$)).mp. | 74543 |
| 16 | or/8-15 | 133338 |
| 17 | exp adolescent/ or exp child/ or exp infant/ or (infant disease* or childhood disease* or adolescen* or babies or baby or boy? or boyfriend or boyhood or child* or girl? or infant* or juvenil* or kid? or minors or minors* or neonat* or neonat* or newborn* or new-born* or paediatric* or peadiatric* or pediatric* or perinat* or preschool* or puber* or pubescen* or school* or teen* or toddler? or underage? or under-age? or youth* or "aged 0-6 years" or "aged 0-5 years" or "aged 0-4 years" or "aged 0-3 years" or "aged 0-2 years" or "aged 0-1 years" or "aged < 6 years" or "aged < 5 years" or "aged < 4 years" or "aged < 3 years" or "aged < 2 years" or "aged < 1 years" or "aged 18 months" or "aged 12 months").mp. | 4912654 |
| 18 | Oral Health/ | 162383 |
| 19 | (Life Quality or Health-Related Quality Of Life or Health Related Quality Of Life).mp. or HRQOL.ab,ti,kw. | 93736 |
| 20 | Dental health/ | 4335 |
| 21 | exp Periodontal Diseases/ or periodontal disease\$.mp. | 110425 |
| 22 | exp Dental Caries/ | 50853 |
| 23 | (carie\$ or carie*).mp. | 61639 |
| 24 | ((teeth or tooth or dental or enamel or dentin or root\$) adj5 (cavit\$ or caries\$ or carious or decay\$ or lesion\$ or deminerali\$ or reminerali\$)).mp. | 73327 |
| 25 | ((teeth or tooth or dental or enamel or dentin) and plaque).mp. | 28386 |
| 26 | exp Tooth demineralization/ | 228225 |
| 27 | Fluorosis, dental/ or (fluorosis or fluorosed or tooth discolouration).mp. | 5521 |
| 28 | Dental enamel/ | 19979 |
| 29 | ("DMF Index" or "Dental Plaque Index").mp. | 1402 |
| 30 | Tooth loss/ or tooth loss.ti,ab,hw,kw. | 35312 |
| 31 | or/18-30 | 510083 |
| 32 | 7 and 16 and 17 and 31 | 498 |

| # | Searches | Results |
|----|--|---------|
| 33 | 32 not ((exp animal/ or nonhuman/) not exp human/) | 492 |
| 34 | 33 not (letter or comment or editorial or newspaper article).pt. | 481 |

7.1.3 Cochrane Central (1946-07 December 2021)

| ID | Search | Hits |
|-----|---|-------|
| #1 | MeSH descriptor: [Fluorides] explode all trees | 2718 |
| #2 | MeSH descriptor: [Fluorine] explode all trees | 85 |
| #3 | MeSH descriptor: [Fluoridation] explode all trees | 39 |
| #4 | ((fluorid* or fluorin* or flurin* or flurid* or flourid* or flourin*)) | 6410 |
| #5 | #1 or #2 or #3 or #4 | 6490 |
| #6 | MeSH descriptor: [Water Supply] explode all trees | 181 |
| #7 | MeSH descriptor: [Water] explode all trees | 2473 |
| #8 | ("water treatment") | 330 |
| #9 | water NEAR fluorid* | 257 |
| #10 | ("community water" OR "community-based water" OR "community supply" OR "community fluoridation") | 30 |
| #11 | #6 OR #7 OR #8 OR #9 OR #10 | 3103 |
| #12 | #5 and #11 | 308 |
| #13 | MeSH descriptor: [Oral Health] explode all trees | 487 |
| #14 | MeSH descriptor: [Tooth Diseases] explode all trees | 11605 |
| #15 | MeSH descriptor: [DMF Index] explode all trees | 519 |
| #16 | MeSH descriptor: [Dental Enamel] explode all trees | 1182 |

| | | |
|-----|--|--------|
| #17 | ("oral health" OR "dental health"):ti,ab,kw | 4084 |
| #18 | (caries OR carious OR cavit* OR decay* OR demineral* OR remineral* OR "dental plaque index") | 23464 |
| #19 | (fluorosis or fluorosed OR ((tooth OR teeth) NEXT (discolour* OR discolor*))) | 797 |
| #20 | (enamel OR root OR dentin OR tooth OR teeth OR oral OR dental):ti,ab,kw | 214033 |
| #21 | (deminerali* in All Text or reminerali* in All Text) | 84 |
| #22 | ("quality of life" OR "life quality" OR QoL OR HRQoL):ti,ab,kw | 128184 |
| #23 | #13 OR #14 OR #15 OR #16 OR #17 OR #18 OR #19 OR #20 OR #21 OR #22 | 336679 |
| #24 | MeSH descriptor: [Child] explode all trees | 59662 |
| #25 | MeSH descriptor: [Pediatrics] explode all trees | 713 |
| #26 | ("early childhood") | 2568 |
| #27 | #24 or #25 or #26 | 61708 |
| #28 | MeSH descriptor: [Fluorides, Topical] explode all trees | 645 |
| #29 | (toothpaste* or tooth-paste* or mouthrins* or mouth-rins* or mouthwash* or mouth-wash* or gel* or varnish* or seal* or paste* or dentifrice* or gum* or lozenge* or drop* or rins*):ti,ab,kw | 78987 |
| #30 | (cariostatic or fluorid\$ or fluor or "PPM F" or PPMF or APF or NAF or "Sodium F" or "Amine F" or SNF2 or "Stannous F" or "phosphate\$ F" or "acidulate\$ F" or "phosphat\$ fluor\$" or | 2226 |

| | | |
|-----|---|------|
| | fluorophosphat\$ or SMFP or MFP or monofluor\$):ti,ab,kw | |
| #31 | MeSH descriptor: [Toothpastes] explode all trees | 849 |
| #32 | #28 #29 or #30 or #31 | 2943 |
| #33 | #12 AND #23 and #27 and #32 | 51 |

7.1.4 Latin American and Caribbean Health Sciences Literature Database (LILACS) (1998-27 July 2021)

| # | Searches | Result |
|--------|--|--------|
| #1 | (fluoride OR fluorine OR fluori\$ OR fluoruro OR fluoreto) AND (water OR public water OR community water) | |
| #2 | ((teeth OR tooth OR dental OR dentin\$ OR enamel OR root\$) OR (Cavit\$ OR caries OR cariou\$ OR decay*)) AND | |
| #3 | (child* OR infant OR baby OR babies OR adolescent OR girl* OR boy* OR paediatric* OR pediatric*) AND (topical OR varnish OR paste OR toothpaste OR gel OR dentifrice OR bath OR rinse) | |
| #4 | type_of_study:("prevalence_studies" OR "evaluation_studies" OR "clinical_trials" OR "prognostic_studies" OR "observational_studies" OR "incidence_studies")) | |
| Total: | | 96 |

7.1.5 Supplementary grey literature table

Scoping searches for Question 2A and 2B were carried out in the search engine *Google.com* to gain an initial idea of terminology and likely key terms relating to the search concepts. Literature and systematic reviews in the area were reviewed to develop search language. Search terms used included combinations of water, fluoridated water, fluoride, oral health, dental health, plus children, and topical fluoride(s). See

Table 2 for the language used in this structured search. Broad terms were used to capture as much relevant material as possible. Further searches were carried out using the websites of relevant bodies. Updated searches of these resources was undertaken in March, 2023, as well as the search engine, *DuckDuckGo*.

Table 29 Grey literature resources for Q 1, 2A and 2B

| Organisation | Website |
|---|---|
| America | |
| American Academy of Oral Medicine (AAOM) | https://www.aaom.com/ |
| American Association of Pediatric Dentistry (AAPD) | https://www.aapd.org/ |
| American Dental Association (ADA) | https://www.ada.org/en |
| Centers for Disease Control (CDC) | https://www.cdc.gov/fluoridation/index.html |
| Department of Health and Human Services | https://www.hhs.gov/ |
| Environmental Protection Agency (EPA) | https://www.epa.gov/ |
| Australia | |
| Australian Dental Association | https://www.ada.org.au/about |
| Department of Health | https://www.health.gov.au/ |
| National Health and Medical Research Council (NHMRC) | https://www.nhmrc.gov.au/ |
| Canada | |
| Canadian Dental Association (CDA) | https://www.cda-adc.ca/en/index.asp |
| Canadian Institute for Health Information (CIHI) | https://www.cihi.ca/en |
| Health Canada | https://www.canada.ca/en/health-canada.html |
| University of Toronto LibGuide (Dentistry/conference proceedings) | https://guides.library.utoronto.ca/c.php?g=250649&p=5001577 |
| Ireland | |
| Dental Council | http://www.dentalcouncil.ie |
| Department of Health | https://www.gov.ie/en/organisation/departments-of-health/ |
| Environmental Protection Agency (EPA) | https://www.epa.ie/ |
| Health Service Executive (HSE) | https://www.hse.ie/eng/ |
| Irish Dental Association | https://www.dentist.ie/ |
| Irish Expert Body on Fluorides and Health | https://tinyurl.com/yntxxdhz |
| New Zealand | |
| Environmental Health Intelligence New Zealand (EHINZ) | https://www.ehinz.ac.nz/ |
| Fluoride Reference Group | https://www.nhmrc.gov.au/about-us/leadership-and-governance/committees/fluoride-reference-group |
| Ministry of Health | https://www.health.govt.nz/ |
| United Kingdom | |
| British Dental Association (BDA) | https://www.bda.org/ |
| Department of Health & Social Care | https://www.gov.uk/government/organisations/department-of-health-and-social-care |
| National Health Service (NHS) | https://www.nhs.uk/ |

| Organisation | Website |
|---|---|
| National Institute for health and care excellence (NICE) | https://www.nice.org.uk/ |
| Scottish dental clinical effectiveness programme | https://www.sdcep.org.uk |
| International Bodies | |
| Centre for Evidence-based Dentistry | https://www.cebd.org/ |
| Council of European Dentists (CED) | https://cedentists.eu/ |
| European Food Safety Authority | https://www.efsa.europa.eu/en |
| International Association for Dental Research (IADR) | https://www.iadr.org/ |
| International Network of Agencies for Health Technology Assessment (INAHTA) | https://database.inahta.org/about#about-inahta |
| World Dental Federation (FDI) | https://www.fdiworlddental.org/ |
| WHO Oral Health Observatory | https://www.who.int/health-topics/oral-health/#tab=tab_1 |
| Search Engines | |
| Google | https://www.google.com/ |
| DuckDuckGo | https://duckduckgo.com/DuckDuckGo?ia=web |
| Research Repositories | |
| Canada's Drug and Health Technology Agency (CADTH) | https://www.cadth.ca/grey-matters-practical-tool-searching-health-related-grey-literature-0 |
| CORE (CONnecting REpositories) open-source repository | https://core.ac.uk/ |
| Health Systems Evidence | https://www.healthsystemsevidence.org/ |
| International Network of Agencies for Health Technology Assessment (INAHTA) | https://database.inahta.org/about#about-inahta |
| Latin American and Caribbean Health Sciences Literature (LILACS) | https://lilacs.bvsalud.org/en/ |
| medRxiv | https://www.medrxiv.org/ |
| RAND | https://www.rand.org/help/search.html |
| Trail Registries | |
| EU Clinical Trials Register | https://www.clinicaltrialsregister.eu/ctr-search/search?query=fluoride+and+oral+health |
| International Clinical Trials Registry Platform (ICTRP) | https://www.who.int/clinical-trials-registry-platform |
| Oral Health Data Portal (World Health Organization) | https://www.who.int/data/gho/data/themes/oral-health-data-portal |

7.2 Appendix B PRISMA checklist and PRISMA-S for Question 2A

7.2.1 PRISMA checklist for Question 2A

| Topic | Item | Checklist item | Location where item is reported |
|-------------------------|------|--|---|
| TITLE | | | |
| Title | 1 | Identify the report as a systematic review. | Title page |
| ABSTRACT | | | |
| Abstract | 2 | See the PRISMA 2020 for Abstracts checklist. | Executive summary |
| INTRODUCTION | | | |
| Rationale | 3 | Describe the rationale for the review in the context of existing knowledge. | Section 1.1.4 |
| Objectives | 4 | Provide an explicit statement of the objective(s) or question(s) the review addresses. | Section 1.2 |
| METHODS | | | |
| Eligibility criteria | 5 | Specify the inclusion and exclusion criteria for the review and how studies were grouped for the syntheses. | Section 2.3.2 |
| Information sources | 6 | Specify all databases, registers, websites, organisations, reference lists and other sources searched or consulted to identify studies. Specify the date when each source was last searched or consulted. | Section 2.4 |
| Search strategy | 7 | Present the full search strategies for all databases, registers and websites, including any filters and limits used. | Appendix A of Section 7, |
| Selection process | 8 | Specify the methods used to decide whether a study met the inclusion criteria of the review, including how many reviewers screened each record and each report retrieved, whether they worked independently, and if applicable, details of automation tools used in the process. | Section 2.6 Error! Reference source not found. |
| Data collection process | 9 | Specify the methods used to collect data from reports, including how many reviewers collected data from each report, whether they worked independently, any processes for obtaining or confirming data from study investigators, and if applicable, details of automation tools used in the process. | Section 2.7 |

| Topic | Item | Checklist item | Location where item is reported |
|-------------------------------|------|---|--|
| Data items | 10a | List and define all outcomes for which data were sought. Specify whether all results that were compatible with each outcome domain in each study were sought (e.g. for all measures, time points, analyses), and if not, the methods used to decide which results to collect. | Section 2.3.2, Table 3, Section 2.7.1.3 |
| | 10b | List and define all other variables for which data were sought (e.g. participant and intervention characteristics, funding sources). Describe any assumptions made about any missing or unclear information. | Section 2.7.1.3 |
| Study risk of bias assessment | 11 | Specify the methods used to assess risk of bias in the included studies, including details of the tool(s) used, how many reviewers assessed each study and whether they worked independently, and if applicable, details of automation tools used in the process. | Section 2.8 |
| Effect measures | 12 | Specify for each outcome the effect measure(s) (e.g. risk ratio, mean difference) used in the synthesis or presentation of results. | Section 2.9.1 |
| Synthesis methods | 13a | Describe the processes used to decide which studies were eligible for each synthesis (e.g. tabulating the study intervention characteristics and comparing against the planned groups for each synthesis (item #5)). | Section 2.9.2 |
| | 13b | Describe any methods required to prepare the data for presentation or synthesis, such as handling of missing summary statistics, or data conversions. | Section 2.9.3 |
| | 13c | Describe any methods used to tabulate or visually display results of individual studies and syntheses. | Section 2.9.2 and 2.9.3 |
| | 13d | Describe any methods used to synthesize results and provide a rationale for the choice(s). If meta-analysis was performed, describe the model(s), method(s) to identify the presence and extent of statistical heterogeneity, and software package(s) used. | Section 2.9.3 |
| | 13e | Describe any methods used to explore possible causes of heterogeneity among study results (e.g. subgroup analysis, meta-regression). | Section 2.9.3 |
| | 13f | Describe any sensitivity analyses conducted to assess robustness of the synthesized results. | Section 2.9.3 |
| Reporting bias assessment | 14 | Describe any methods used to assess risk of bias due to missing results in a synthesis (arising from reporting biases). | Section 2.9.2 and 2.9.3 |

| Topic | Item | Checklist item | Location where item is reported |
|-------------------------------|------|--|---|
| Certainty assessment | 15 | Describe any methods used to assess certainty (or confidence) in the body of evidence for an outcome. | Section 2.10 |
| RESULTS | | | |
| Study selection | 16a | Describe the results of the search and selection process, from the number of records identified in the search to the number of studies included in the review, ideally using a flow diagram. | Section 3.2.1, Appendix F of Section 7 |
| | 16b | Cite studies that might appear to meet the inclusion criteria, but which were excluded, and explain why they were excluded. | Appendix C of Section 7.3 |
| Study characteristics | 17 | Cite each included study and present its characteristics. | Section 3.2.2, Table 45 |
| Risk of bias in studies | 18 | Present assessments of risk of bias [and/or quality assessment] for each included study. | Appendix H of Section 7 |
| Results of individual studies | 19 | For all outcomes, present, for each study: (a) summary statistics for each group (where appropriate) and (b) an effect estimate and its precision (e.g. confidence/credible interval), ideally using structured tables or plots. | Sections 3.2.4 |
| Results of syntheses | 20a | For each synthesis, briefly summarise the characteristics and risk of bias among contributing studies. | Sections 3.2.3, Table 46, Appendix H of Section 7 |
| | 20b | Present results of all statistical syntheses conducted. If meta-analysis was done, present for each the summary estimate and its precision (e.g. confidence/credible interval) and measures of statistical heterogeneity. If comparing groups, describe the direction of the effect. | Section 3.2.4.2.18 |
| | 20c | Present results of all investigations of possible causes of heterogeneity among study results. | Not applicable |
| | 20d | Present results of all sensitivity analyses conducted to assess the robustness of the synthesized results. | Not applicable |
| Reporting biases | 21 | Present assessments of risk of bias due to missing results (arising from reporting biases) for each synthesis assessed. | Not applicable as mainly cross section surveys |

| Topic | Item | Checklist item | Location where item is reported |
|--|------|--|---|
| Certainty of evidence | 22 | Present assessments of certainty (or confidence) in the body of evidence for each outcome assessed. | Section 3.2.4.2.20 |
| DISCUSSION | | | |
| Discussion | 23a | Provide a general interpretation of the results in the context of other evidence. | Sections 4.1 and 4.2 |
| | 23b | Discuss any limitations of the evidence included in the review. | Section 4.3 |
| | 23c | Discuss any limitations of the review processes used. | Section 4.3 |
| | 23d | Discuss implications of the results for practice, policy, and future research. | Sections 4.4 and 4.5 |
| OTHER INFORMATION | | | |
| Registration and protocol | 24a | Provide registration information for the review, including register name and registration number, or state that the review was not registered. | Section 2.3.2 |
| | 24b | Indicate where the review protocol can be accessed, or state that a protocol was not prepared. | Section 2.3.2 |
| | 24c | Describe and explain any amendments to information provided at registration or in the protocol. | Section 2.3.2 |
| Support | 25 | Describe sources of financial or non-financial support for the review, and the role of the funders or sponsors in the review. | Not applicable as all authors are salaried public servants who are funded from the DOH public funding and are obliged to be objective |
| Competing interests | 26 | Declare any competing interests of review authors. | None |
| Availability of data, code and other materials | 27 | Report which of the following are publicly available and where they can be found: template data collection forms; data extracted from included studies; data used for all analyses; analytic code; any other materials used in the review. | Appendix D and E of Section 7 |

Source: Page *et al.* (2021)[1]

7.2.2 PRISMA-S Question 2A

| Section/ topic | # | Checklist item | Location(s) Reported |
|--|----|--|--|
| INFORMATION SOURCES AND METHODS | | | |
| Database name | 1 | Name each individual database searched, stating the platform for each. | Sections 2.4.4 and 2.5 |
| Multi-database searching | 2 | If databases were searched simultaneously on a single platform, state the name of the platform, listing all of the databases searched. | n/a |
| Study registries | 3 | List any study registries searched. | Sections 2.4 and 2.5 |
| Online resources and browsing | 4 | Describe any online or print source purposefully searched or browsed (e.g., tables of contents, print conference proceedings, web sites), and how this was done. | Sections 2.4.5 and 2.4.6 |
| Citation searching | 5 | Indicate whether cited references or citing references were examined, and describe any methods used for locating cited/citing references (e.g., browsing reference lists, using a citation index, setting up email alerts for references citing included studies). | Section 2.4.5 |
| Contacts | 6 | Indicate whether additional studies or data were sought by contacting authors, experts, manufacturers, or others. | n/a |
| Other methods | 7 | Describe any additional information sources or search methods used. | n/a |
| SEARCH STRATEGIES | | | |
| Full search strategies | 8 | Include the search strategies for each database and information source, copied and pasted exactly as run. | Appendix A of Section 7 |
| Limits and restrictions | 9 | Specify that no limits were used, or describe any limits or restrictions applied to a search (e.g., date or time period, language, study design) and provide justification for their use. | Sections 2.5 and Appendix A of Section 7 |
| Search filters | 10 | Indicate whether published search filters were used (as originally designed or modified), and if so, cite the filter(s) used. | n/a |
| Prior work | 11 | Indicate when search strategies from other literature reviews were adapted or reused for a substantive part or all of the search, citing the previous review(s). | n/a |

| | | | |
|-------------------------|--------|--|-------------------------|
| Updates | 1 2 | Report the methods used to update the search(es) (e.g., rerunning searches, email alerts). | Appendix A of Section 7 |
| Dates of searches | 1 3 | For each search strategy, provide the date when the last search occurred. | Appendix A of Section 7 |
| PEER REVIEW | | | |
| Peer review | 1 4 | Describe any search peer review process. | Section 2.4.3 |
| MANAGING RECORDS | | | |
| Total Records | 1 5 | Document the total number of records identified from each database and other information sources. | Appendix A of Section 7 |
| Deduplication | 1 6 | Describe the processes and any software used to deduplicate records from multiple database searches and other information sources. | Section 2.4.4 |

PRISMA-S: An Extension to the PRISMA Statement for Reporting Literature Searches in Systematic Reviews

Rethlefsen ML, Kirtley S, Waffenschmidt S, Ayala AP, Moher D, Page MJ, Koffel JB, PRISMA-S Group.

Last updated February 27, 2020.

7.3 Appendix C Studies excluded at full text and extraction screening stages

7.3.1 Exclude on population

Exclude on population n= 8

Aimee NR, van Wijk AJ, Maltz M, *et al.* Dental caries, fluorosis, oral health determinants, and quality of life in adolescents. *Clin Oral Investig* 2017;21:1811–20. doi:10.1007/s00784-016-1964-3

Cabral RN, Leal SC, Bernardino Í de M, *et al.* Caries lesion transition patterns of schoolchildren in a fluoridated community in Brazil. *Clin Oral Investig* 2022;26:689–95. doi:10.1007/s00784-021-04046-9

Chattopadhyay A, Arevalo O, Cecil JC. Kentucky's oral health indicators and progress towards Healthy People 2010 objectives. *J Ky Med Assoc* 2008;106:165–74.

Devoto FC, Bordoni NE, De Manfredi CF. Dental caries in deciduous teeth of nineteenth century Araucanians. *J Dent Res* 1968;47:571–4.

Hausen H, Heinonen OP, Paunio I. Fluoride exposure combinations and caries in permanent dentition among Finnish children. *Community Dent Oral Epidemiol* 1981;9:108–11. doi:10.1111/j.1600-0528.1981.tb01039.x

Lawrence H P, Sheiham A. Caries progression in 12- to 16-year-old schoolchildren in fluoridated and fluoride-deficient areas in Brazil. *Community Dent Oral Epidemiol* 1997;25:402–11.

Maupome G, Shulman JD, Clark DC, *et al.* Socio-demographic features and fluoride technologies contributing to higher fluorosis scores in permanent teeth of Canadian children. *Caries Res* 2003;37:327–34. doi:10.1159/000072163

Peterson JK. A supervised brushing trial of sodium monofluorophosphate dentrifices in a fluoridated area. *Caries Res* 1979;13:68–72. doi:10.1159/000260385

7.3.2 Exclude on intervention

Exclude on intervention (n=149)

Adair SM, Hanes CM, Russell CM, *et al.* Dental caries and fluorosis among children in a rural Georgia area. *Pediatr Dent* 1999;21:81–5.

Agus HM, Schamschula RG, Barmes DE, *et al.* Associations between the total fluoride content of dental plaque and individual caries experience in Australian children. *Community Dent Oral Epidemiol* 1976;4:210–4. doi:10.1111/j.1600-0528.1976.tb00986.x

Ahmed AT, Soto-Rojas A, Dean J, *et al.* Demarcated Primary Second Molar Hypomineralization: Prevalence Data and Associated Sociodemographic Determinants from Indiana. *Pediatr Dent* 2021;43:443–50.

Ahmed NAM, Astrøm AN, Skaug N, *et al.* Dental caries prevalence and risk factors among 12-year old schoolchildren from Baghdad, Iraq: a post-war survey. *Int Dent J* 2007;57:36–44. doi:10.1111/j.1875-595x.2007.tb00116.x

Ainamo J, Parviainen K. Occurrence of plaque, gingivitis and caries as related to self reported frequency of toothbrushing in fluoride areas in Finland. *Community Dent Oral Epidemiol* 1979;7:142–6. doi:10.1111/j.1600-0528.1979.tb01202.x

Ainamo J, Parviainen K. Influence of increased toothbrushing frequency on dental health in low, optimal, and high fluoride areas in Finland. *Community Dent Oral Epidemiol* 1989;17:296–9. doi:10.1111/j.1600-0528.1989.tb00640.x

Aleksejuniene J, Arneberg P, Eriksen HM. Caries prevalence and oral hygiene in Lithuanian children and adolescents. *Acta Odontol Scand* 1996;54:75–80. doi:10.3109/00016359609003513

Exclude on intervention (n=149)

Armfield JM, Spencer AJ, Roberts-Thomson KF, *et al.* Water fluoridation and the association of sugar-sweetened beverage consumption and dental caries in Australian children. *Am J Public Health* 2013;103:494–500. doi:10.2105/AJPH.2012.300889

Arra MC, Lemke C. EFFECTS OF ADJUSTED FLUORIDATED WATER ON DENTAL CARIES IN SCHOOL CHILDREN OF AMERY, WIS. *J Am Dent Assoc* 1964;69:460–4. doi:10.14219/jada.archive.1964.0317

Arra MC, Lemke C. REDUCTION OF DENTAL CARIES IN CHILDREN THROUGH ADJUSTED FLUORIDATED WATER. *N Y J Dent* 1965;35:57 PASSIM.

Ashkenazi M, Bidoosi M, Levin L. Effect of Preventive Oral Hygiene Measures on the Development of New Carious lesions. *Oral Health Prev Dent* 2014;12:61–9. doi:10.3290/j.ohpd.a31219

Ashkenazi M, Cohen R, Levin L. Self-reported compliance with preventive measures among regularly attending pediatric patients. *J Dent Educ* 2007;71:287–95.

Australian Research Centre for Population Oral Health, The University of Adelaide, South Australia. The benefits of water fluoridation across areas of differing socio-economic status. *Aust Dent J* 2008;53:180–3. doi:10.1111/j.1834-7819.2008.00030.x

Bagramian RA. A 5-year school-based comprehensive preventive program in Michigan, U.S.A. *Community Dent Oral Epidemiol* 1982;10:234–7. doi:10.1111/j.1600-0528.1982.tb00385.x

Bagramian RA, Graves RC, Bhat M. A combined approach to preventing dental caries in schoolchildren: caries reductions after one year. *J Am Dent Assoc* 1976;93:1014–9. doi:10.14219/jada.archive.1976.0032

Barnes VM, Santarpia P, Richter R, *et al.* Clinical evaluation of the anti-plaque effect of a commercial chewing gum. *J Clin Dent* 2005;16:1–5.

Barnhart WE, Hiller LK, Leonard GJ, *et al.* Dentifrice usage and ingestion among four age groups. *J Dent Res* 1974;53:1317–22. doi:10.1177/00220345740530060301

Barrett MJ, Williamson JJ. Oral health of Australian aborigines: survey methods and prevalence of dental caries. *Aust Dent J* 1972;17:37–50. doi:10.1111/j.1834-7819.1972.tb02744.x

Beal JF, James PM. Dental caries prevalence in 5-year-old children following five and a half years of water fluoridation in Birmingham. *Br Dent J* 1971;130:284–8. doi:10.1038/sj.bdj.4802658

Beltrán-Aguilar ED, Barker L, Dye BA. Prevalence and severity of dental fluorosis in the United States, 1999–2004. *NCHS Data Brief* 2010;;1–8.

Beltrán-Aguilar ED, Griffin SO, Lockwood SA. Prevalence and trends in enamel fluorosis in the United States from the 1930s to the 1980s. *J Am Dent Assoc* 2002;133:157–65. doi:10.14219/jada.archive.2002.0139

Biesbrock AR, Gerlach RW, Bollmer BW, *et al.* Relative anti-caries efficacy of 1100, 1700, 2200, and 2800 ppm fluoride ion in a sodium fluoride dentifrice over 1 year. *Community Dent Oral Epidemiol* 2001;29:382–9. doi:10.1034/j.1600-0528.2001.290508.x

Bixler D, Muhler JC. COMBINED USE OF THREE AGENTS CONTAINING STANNOUS FLUORIDE: A PROPHYLACTIC PASTE, A SOLUTION AND A DENTIFRICE. *J Am Dent Assoc* 1964;68:792–800. doi:10.14219/jada.archive.1964.0195

Bixler D, Muhler JC. Effectiveness of a stannous fluoride-containing dentifrice in reducing dental caries in children in a boarding school environment. *J Am Dent Assoc* 1966;72:653–8. doi:10.14219/jada.archive.1966.0076

Blinkhorn AS, Byun R, Mehta P, *et al.* A 4-year assessment of a new water-fluoridation scheme in New South Wales, Australia. *Int Dent J* 2015;65:156–63. doi:10.1111/idj.12166

Bohaty BS, Parker WA, Seale NS, *et al.* The prevalence of fluorosis-like lesions associated with topical and systemic fluoride usage in an area of optimal water fluoridation. *Pediatr Dent* 1989;11:125–8.

Exclude on intervention (n=149)

Bonow MLM, Azevedo MS, Goettens ML, *et al.* Efficacy of 1.23% APF gel applications on incipient carious lesions: a double-blind randomized clinical trial. *Braz Oral Res* 2013;27:279–85.

doi:10.1590/S1806-83242013000300007

Bottenberg P, Declerck D, Ghidry W, *et al.* Prevalence and determinants of enamel fluorosis in Flemish schoolchildren. *Caries Res* 2004;38:20–8. doi:10.1159/000073916

Bramlett MD, Soobader M-J, Fisher-Owens SA, *et al.* Assessing a multilevel model of young children's oral health with national survey data. *Community Dent Oral Epidemiol* 2010;38:287–98.

doi:10.1111/j.1600-0528.2010.00536.x

Brodeur JM, Simard PL, Demers M, *et al.* Comparative effects of FMR programs in fluoridated and unfluoridated communities. *J Can Dent Assoc* 1988;54:761–5.

Broffitt B, Levy SM, Warren J, *et al.* Factors associated with surface-level caries incidence in children aged 9 to 13: the Iowa Fluoride Study. *J Public Health Dent* 2013;73:304–10. doi:10.1111/jphd.12028

Bronson ME. Dental health in an area of maximum water fluoridation. *Dent Hyg (Chic)* 1982;56:38–41.

Broughton JR, Person M, Maipi JTH, *et al.* Ukaipō niho: the place of nurturing for oral health. *N Z Dent J* 2014;110:18–23.

Brunelle JA, Carlos JP. Recent trends in dental caries in U.S. children and the effect of water fluoridation. *J Dent Res* 1990;69 Spec No:723–7; discussion 820–823.

doi:10.1177/00220345900690S141

Burton VJ, Rob MI, Craig GG, *et al.* Changes in the caries experience of 12-year-old Sydney schoolchildren between 1963 and 1982. *Med J Aust* 1984;140:405–7. doi:10.5694/j.1326-5377.1984.tb108100.x

Butler WJ, Segreto V, Collins E. Prevalence of dental mottling in school-aged lifetime residents of 16 Texas communities. *Am J Public Health* 1985;75:1408–12. doi:10.2105/ajph.75.12.1408

Chankanka O, Cavanaugh JE, Levy SM, *et al.* Longitudinal associations between children's dental caries and risk factors. *J Public Health Dent* 2011;71:289–300. doi:10.1111/j.1752-7325.2011.00271.x

Choo-Wosoba H, Levy SM, Datta S. Marginal regression models for clustered count data based on zero-inflated Conway-Maxwell-Poisson distribution with applications. *Biometrics* 2016;72:606–18.

doi:10.1111/biom.12436

Colquhoun J. Influence of social class and fluoridation on child dental health. *Community Dent Oral Epidemiol* 1985;13:37–41. doi:10.1111/j.1600-0528.1985.tb00417.x

Cons NC, Janerich DT, Senning RS. Albany topical fluoride study. *J Am Dent Assoc* 1970;80:777–81. doi:10.14219/jada.archive.1970.0113

Creedon MI, O'Mullane DM. Factors affecting caries levels amongst 5-year-old children in County Kerry, Ireland. *Community Dent Health* 2001;18:72–8.

de Sousa M da LR, Wagner M, Sheiham A. Caries reductions related to the use of fluorides: a retrospective cohort study. *Int Dent J* 2002;52:315–20. doi:10.1002/j.1875-595x.2002.tb00877.x

Dean HT, Jay P, ARNOLD FA Jr, *et al.* Nutrition classics. Public Health Reports, Volume 56, 1941, pages 761–792. Domestic water and dental caries. II. A study of 2,832 white children, aged 12–14 years, of 8 suburban Chicago communities, including *Lactobacillus acidophilus* studies of 1,761 children. By H. Trendley Dean, Philip Jay, Francis A. Arnold, Jr., and Elias Elvove. *Nutr Rev* 1976;34:116–8.

doi:10.1111/j.1753-4887.1976.tb05724.x

Dimitropoulos Y, Holden A, Gwynne K, *et al.* Outcomes of a co-designed, community-led oral health promotion program for Aboriginal children in rural and remote communities in New South Wales, Australia. *Community Dent Health* 2020;37:132–7. doi:10.1922/CDH_00005Dimitropoulos06

Do LG, Spencer AJ. Decline in the prevalence of dental fluorosis among South Australian children. *Community Dent Oral Epidemiol* 2007;35:282–91. doi:10.1111/j.1600-0528.2007.00314.x

Exclude on intervention (n=149)

Do LG, Spencer AJ. Risk-benefit balance in the use of fluoride among young children. *J Dent Res* 2007;86:723–8. doi:10.1177/154405910708600807

do Nascimento HAR, Soares Ferreira JM, Granville-Garcia AF, *et al.* Estimation of toothpaste fluoride intake in preschool children. *Braz Dent J* 2013;24:142–6. doi:10.1590/0103-6440201302087

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Exclude on comparator n=1

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7.4 Appendix D Extraction form

Table 30 General information data form

| Study ID From Eppi | Author First or | Year Of publication | Location County | Area State/County/City/Town | Objective Aim of study | Secondary publication Data will not be extracted unless additional endpoints | Associated papers Same overall project differences | Study design HRB decision | Participant age Mean or ranges described in study | Artificial fluoridation Confirm if explicitly stated (Y/N) | Fluoride interventions | Outcome Oral health outcome assessed | Outcome details Including method of measurement | Extracted | Validated |
|--------------------|-----------------|---------------------|-----------------|-----------------------------|------------------------|--|--|---------------------------|---|--|------------------------|--------------------------------------|---|-----------|-----------|
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Table 31 Study design data form

| Study ID | Author | Year | Study design (Author allocated) | Study design (HRB allocated) | Justification | Length of study | Length of exposure to CWF | Details of exposure | Details of comparator | Eligibility criteria | Sample size calculation | Response rate | Blinding of assessors to exposure | % Lost to follow-up | Method for handling missing data | Data collection | Confounders | Control for confounding | Identification of effect modification | Effect modifiers | Notes |
|----------|--------------|----------------|---------------------------------|------------------------------|---------------|-----------------|--|---------------------|-----------------------|----------------------|--|---------------|-----------------------------------|---------------------|---------------------------------------|-------------------|-------------|-------------------------|---------------------------------------|------------------|-------|
| Form Epi | First author | Of publication | As stated in the study | As agreed by research team | | | Length of time exposed to community water fluoridation | Including dose | Including dose | | expected prevalence, power to detect a difference and allowed variance, results CIs calculated | | | For main analysis | e.g. last observation carried forward | Brief description | | | Yes or not reported | | |
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Table 32 Study participants data form

| Study ID | Author | Year | Group for characteristics | N | Mean age/Age range | % Female | N included in final analysis |
|------------------|---------------------|-----------------------|---------------------------|-----------------|--------------------|----------|------------------------------|
| <i>From Eppi</i> | <i>First author</i> | <i>Of publication</i> | | <i>Enrolled</i> | | | |
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Table 33 Outcomes

| Of publica tion | Outco me of intere st: Caries | % carie s prim ary teet h | % carie s free prim ary teet h | % caries perma nent teeth | % caries free perma nent teeth | dmft/ deft | dmfs/ defs | DM FT | DM FS | Method of caries identific ation | Clinical examin ation criteria | Outco me of intere st: Fluor osis | Fluor osis (Dea n's index) | Fluoro sis (Thylst rup- Fejersk ov index) | Tooth Surfa ces Index of Fluor osis | Type of teeth exami ned for fluoro sis | Hypomineral isation by photographs |
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Table 34 Caries outcome data form using example of primary dentition dmft

| Country | Author | Year | Age in years | CWF ppm | Baseline dmft CWF | Baseline CWF SD | Baseline CWF Total | Final dmft CWF | Final dmft SD CWF | Final CWF Total | Fluoride deficient ppm | Baseline mean dmft No F | Baseline SD No F | Baseline CWF Total | Final dmft No F | Final SD No F | Final No F Total | Difference in % point or dmft |
|---------|--------|------|--------------|---------|-------------------|-----------------|--------------------|----------------|-------------------|-----------------|------------------------|-------------------------|------------------|--------------------|-----------------|---------------|------------------|-------------------------------|
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This table was repeated for dmfs, % with CDC, and % without CDC for primary dentition. The table was also repeated for DMFT, DFMS, % with CDC, and % without CDC for permanent dentition

Table 35 Fluorosis outcome data form

| Country | Author | Year | Age in years | CWF ppm | Baseline % fluorosis | Baseline 95% CI | Baseline CWF affected number | Baseline CWF Total | Final % fluorosis | Final 95% CI | Final CWF affected number | Final CWF Total | Baseline 5 fluorosis No F | Baseline 95% CI No F | Baseline affected number No F | Baseline CWF Total | Final % No F | Final 95%CI No F | Final affected number No F | Final No F Total | Difference in % point or dmft |
|---------|--------|------|--------------|---------|----------------------|-----------------|------------------------------|--------------------|-------------------|--------------|---------------------------|-----------------|---------------------------|----------------------|-------------------------------|--------------------|--------------|------------------|----------------------------|------------------|-------------------------------|
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7.5 Appendix E Quality assessment tools

See 6.5 in Section 6.5 National Heart, Lung, and Blood Institute's (NHLBI's) quality assessment tool for observational cohort studies and cross-sectional survey

Table 36 NHLBI's quality assessment tool for observational case-control studies for question 2A

| Questions NHLBI's quality assessment tool for observational case-control studies | Yes | No | Other*cannot decide, not reported, not applicable |
|---|-----|----|---|
| 1. Was the research question or objective in this paper clearly stated and appropriate? | | | |
| <p>Did the authors describe their goal in conducting this research? Is it easy to understand what they were looking to find? This issue is important for any scientific paper of any type. High quality scientific research explicitly defines a research question.</p> | | | |
| 2. Was the study population clearly specified and defined? | | | |
| <p>Did the authors describe the group of individuals from which the cases and controls were selected or recruited, while using demographics, location, and time period? If the investigators conducted this study again, would they know exactly who to recruit, from where, and from what time period?</p> <p>Investigators identify case-control study populations by location, time period, and inclusion criteria for cases (individuals with the disease, condition, or problem) and controls (individuals without the disease, condition, or problem). For example, the population for a study of lung cancer and chemical exposure would be all incident cases of lung cancer diagnosed in patients ages 35 to 79, from January 1, 2003 to December 31, 2008, living in Texas during that entire time period, as well as controls without lung cancer recruited from the same population during the same time period. The population is clearly described as: (1) who (men and women ages 35 to 79 with (cases) and without (controls) incident lung cancer); (2) where (living in Texas); and (3) when (between January 1, 2003 and December 31, 2008).</p> <p>Other studies may use disease registries or data from cohort studies to identify cases. In these cases, the populations are individuals who live in the area covered by the disease registry or included in a cohort study (i.e., nested case-control or case-cohort). For example, a study of the relationship between vitamin D intake and myocardial infarction might use patients identified via the GRACE registry, a database of heart attack patients.</p> <p>NHLBI staff encouraged reviewers to examine prior papers on methods (listed in the reference list) to make this assessment, if necessary.</p> | | | |

| Questions NHLBI's quality assessment tool for observational case-control studies | Yes | No | Other* cannot decide, not reported, not applicable |
|--|-----|----|--|
|--|-----|----|--|

In order for a study to truly address the research question, the target population—the population from which the study population is drawn and to which study results are believed to apply—should be carefully defined. Some authors may compare characteristics of the study cases to characteristics of cases in the target population, either in text or in a table. When study cases are shown to be representative of cases in the appropriate target population, it increases the likelihood that the study was well-designed per the research question.

However, because these statistics are frequently difficult or impossible to measure, publications should not be penalized if case representation is not shown. For most papers, the response to question 3 will be "NR." Those subquestions are combined because the answer to the second subquestion—case representation—determines the response to this item. However, it cannot be determined without considering the response to the first subquestion. For example, if the answer to the first subquestion is "yes," and the second, "CD," then the response for item 3 is "CD."

3. Did the authors include a sample size justification?

Did the authors discuss their reasons for selecting or recruiting the number of individuals included? Did they discuss the statistical power of the study and provide a sample size calculation to ensure that the study is adequately powered to detect an association (if one exists)? This question does not refer to a description of the manner in which different groups were included or excluded using the inclusion/exclusion criteria (e.g., "Final study size was 1,378 participants after exclusion of 461 patients with missing data" is not considered a sample size justification for the purposes of this question).

An article's methods section usually contains information on sample size and the size needed to detect differences in exposures and on statistical power.

4. Were controls selected or recruited from the same or similar population that gave rise to the cases (including the same timeframe)?

To determine whether cases and controls were recruited from the same population, one can ask hypothetically, "If a control was to develop the outcome of interest (the condition that was used to select cases), would that person have been eligible to become a case?" Case-control studies begin with the selection of the cases (those with the outcome of interest, e.g., lung cancer) and controls (those in whom the outcome is absent). Cases and controls are then evaluated and categorized by their exposure status. For the lung cancer example, cases and controls were recruited from hospitals in a given region. One may reasonably assume that controls in the catchment area for the hospitals, or those already in the hospitals for a different reason, would attend those hospitals if they became a case; therefore, the controls are drawn from the same population as the cases.

| Questions NHLBI's quality assessment tool for observational case-control studies | Yes | No | Other* cannot decide, not reported, not applicable |
|--|-----|----|--|
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If the controls were recruited or selected from a different region (e.g., a State other than Texas) or time period (e.g., 1991-2000), then the cases and controls were recruited from different populations, and the answer to this question would be "no."

The following example further explores selection of controls. In a study, eligible cases were men and women, ages 18 to 39, who were diagnosed with atherosclerosis at hospitals in Perth, Australia, between July 1, 2000 and December 31, 2007. Appropriate controls for these cases might be sampled using voter registration information for men and women ages 18 to 39, living in Perth (population-based controls); they also could be sampled from patients without atherosclerosis at the same hospitals (hospital-based controls). As long as the controls are individuals who would have been eligible to be included in the study as cases (if they had been diagnosed with atherosclerosis), then the controls were selected appropriately from the same source population as cases.

In a prospective case-control study, investigators may enroll individuals as cases at the time they are found to have the outcome of interest; the number of cases usually increases as time progresses. At this same time, they may recruit or select controls from the population without the outcome of interest. One way to identify or recruit cases is through a surveillance system. In turn, investigators can select controls from the population covered by that system. This is an example of population-based controls. Investigators also may identify and select cases from a cohort study population and identify controls from outcome-free individuals in the same cohort study. This is known as a nested case-control study.

5. Were the definitions, inclusion and exclusion criteria, algorithms or processes used to identify or select cases and controls valid, reliable, and implemented consistently across all study participants?

Were the inclusion and exclusion criteria developed prior to recruitment or selection of the study population? Were the same underlying criteria used for all of the groups involved? To answer this question, reviewers determined if the investigators developed I/E criteria prior to recruitment or selection of the study population and if they used the same underlying criteria for all groups. The investigators should have used the same selection criteria, except for study participants who had the disease or condition, which would be different for cases and controls by definition. Therefore, the investigators use the same age (or age range), gender, race, and other characteristics to select cases and controls. Information on this topic is usually found in a paper's section on the description of the study population.

6. Were the cases clearly defined and differentiated from controls?

| Questions NHLBI's quality assessment tool for observational case-control studies | Yes | No | Other* cannot decide, not reported, not applicable |
|--|-----|----|--|
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For this question, reviewers looked for descriptions of the validity of case and control definitions and processes or tools used to identify study participants as such. Was a specific description of "case" and "control" provided? Is there a discussion of the validity of the case and control definitions and the processes or tools used to identify study participants as such? They determined if the tools or methods were accurate, reliable, and objective. For example, cases might be identified as "adult patients admitted to a VA hospital from January 1, 2000 to December 31, 2009, with an ICD-9 discharge diagnosis code of acute myocardial infarction and at least one of the two confirmatory findings in their medical records: at least 2mm of ST elevation changes in two or more ECG leads and an elevated troponin level. Investigators might also use ICD-9 or CPT codes to identify patients. All cases should be identified using the same methods. Unless the distinction between cases and controls is accurate and reliable, investigators cannot use study results to draw valid conclusions.

7. If less than 100 percent of eligible cases and/or controls were selected for the study, were the cases and/or controls randomly selected from those eligible?

If a case-control study did not use 100 percent of eligible cases and/or controls (e.g., not all disease-free participants were included as controls), did the authors indicate that random sampling was used to select controls? When it is possible to identify the source population fairly explicitly (e.g., in a nested case-control study, or in a registry-based study), then random sampling of controls is preferred. When investigators used consecutive sampling, which is frequently done for cases in prospective studies, then study participants are not considered randomly selected. In this case, the reviewers would answer "no" to Question 8. However, this would not be considered a fatal flaw.

If investigators included all eligible cases and controls as study participants, then reviewers marked "NA" in the tool. If 100 percent of cases were included (e.g., NA for cases) but only 50 percent of eligible controls, then the response would be "yes" if the controls were randomly selected, and "no" if they were not. If this cannot be determined, the appropriate response is "CD."

8. Was there use of concurrent controls?

A concurrent control is a control selected at the time another person became a case, usually on the same day. This means that one or more controls are recruited or selected from the population without the outcome of interest at the time a case is diagnosed. Investigators can use this method in both prospective case-control studies and retrospective case-control studies. For example, in a retrospective study of adenocarcinoma of the colon using data from hospital records, if hospital records indicate that Person A was diagnosed with adenocarcinoma of the colon on June 22, 2002, then investigators would select one or more controls from the population of patients without adenocarcinoma of the

| Questions NHLBI's quality assessment tool for observational case-control studies | Yes | No | Other* cannot decide, not reported, not applicable |
|--|-----|----|--|
|--|-----|----|--|

colon on that same day. This assumes they conducted the study retrospectively, using data from hospital records. The investigators could have also conducted this study using patient records from a cohort study, in which case it would be a nested case-control study.

Investigators can use concurrent controls in the presence or absence of matching and vice versa. A study that uses matching does not necessarily mean that concurrent controls were used.

9. Were the investigators able to confirm that the exposure/risk occurred prior to the development of the condition or event that defined a participant as a case?

Investigators first determine case or control status (based on presence or absence of outcome of interest), and then assess exposure history of the case or control; therefore, reviewers ascertained that the exposure preceded the outcome. For example, if the investigators used tissue samples to determine exposure, did they collect them from patients prior to their diagnosis? If hospital records were used, did investigators verify that the date a patient was exposed (e.g., received medication for atherosclerosis) occurred prior to the date they became a case (e.g., was diagnosed with type 2 diabetes)? For an association between an exposure and an outcome to be considered causal, the exposure must have occurred prior to the outcome.

10. Were the measures of exposure/risk clearly defined, valid, reliable, and implemented consistently (including the same time period) across all study participants?

Were the exposure measures defined in detail? Were the tools or methods used to measure exposure accurate and reliable—for example, have they been validated or are they objective? This is important, as it influences confidence in the reported exposures. Equally important is whether the exposures were assessed in the same manner within groups and between groups. This question pertains to bias resulting from exposure misclassification (i.e., exposure ascertainment).

For example, a retrospective self-report of dietary salt intake is not as valid and reliable as prospectively using a standardized dietary log plus testing participants' urine for sodium content because participants' retrospective recall of dietary salt intake may be inaccurate and result in misclassification of exposure status. Similarly, BP results from practices that use an established protocol for measuring BP would be considered more valid and reliable than results from practices that did not use standard protocols. A protocol may include using trained BP assessors, standardized equipment (e.g., the same BP device which has been tested and calibrated), and a standardized procedure (e.g., patient is seated for 5

| Questions NHLBI's quality assessment tool for observational case-control studies | Yes | No | Other* cannot decide, not reported, not applicable |
|--|-----|----|--|
|--|-----|----|--|

minutes with feet flat on the floor, BP is taken twice in each arm, and all four measurements are averaged).

11. Were the assessors of exposure/risk blinded to the case or control status of participants?

Blinding or masking means that outcome assessors did not know whether participants were exposed or unexposed. To answer this question, reviewers examined articles for evidence that the outcome assessor(s) was masked to the exposure status of the research participants. An outcome assessor, for example, may examine medical records to determine the outcomes that occurred in the exposed and comparison groups. Sometimes the person measuring the exposure is the same person conducting the outcome assessment. In this case, the outcome assessor would most likely not be blinded to exposure status. A reviewer would note such a finding in the comments section of the assessment tool.

One way to ensure good blinding of exposure assessment is to have a separate committee, whose members have no information about the study participants' status as cases or controls, review research participants' records. To help answer the question above, reviewers determined if it was likely that the outcome assessor knew whether the study participant was a case or control. If it was unlikely, then the reviewers marked "no" to Question 12. Outcome assessors who used medical records to assess exposure should not have been directly involved in the study participants' care, since they probably would have known about their patients' conditions. If the medical records contained information on the patient's condition that identified him/her as a case (which is likely), that information would have had to be removed before the exposure assessors reviewed the records.

If blinding was not possible, which sometimes happens, the reviewers marked "NA" in the assessment tool and explained the potential for bias.

12. Were key potential confounding variables measured and adjusted statistically in the analyses? If matching was used, did the investigators account for matching during study analysis?

Were key potential confounding variables measured and adjusted for, such as by statistical adjustment for baseline differences? Investigators often use logistic regression or other regression methods to account for the influence of variables not of interest.

This is a key issue in case-controlled studies; statistical analyses need to control for potential confounders, in contrast to RCTs in which the randomization process controls for potential confounders. In the analysis, investigators need to control for all key factors that may be associated

| Questions NHLBI's quality assessment tool for observational case-control studies | Yes | No | Other* cannot decide, not reported, not applicable |
|--|-----|----|--|
|--|-----|----|--|

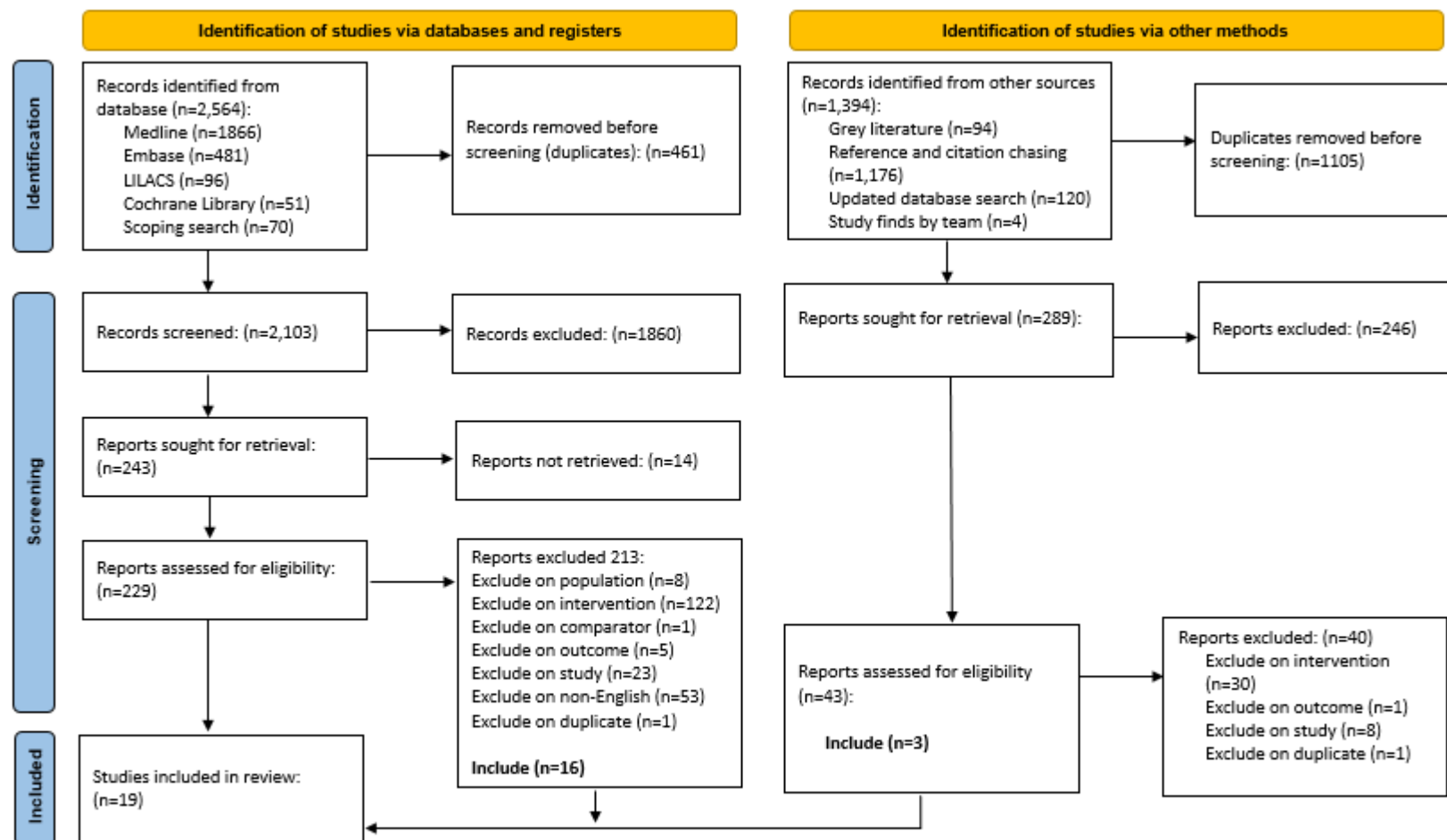
with both the exposure of interest and the outcome and are not of interest to the research question.

A study of the relationship between smoking and CVD events illustrates this point. Such a study needs to control for age, gender, and body weight; all are associated with smoking and CVD events. Well-done case-control studies control for multiple potential confounders.

Matching is a technique used to improve study efficiency and control for known confounders. For example, in the study of smoking and CVD events, an investigator might identify cases that have had a heart attack or stroke and then select controls of similar age, gender, and body weight to the cases. For case-control studies, it is important that if matching was performed during the selection or recruitment process, the variables used as matching criteria (e.g., age, gender, race) should be controlled for in the analysis.

7.6 Appendix F PRISMA flow diagram for Question 2A

PRISMA 2020 flow diagram for new systematic reviews which included searches of databases, registers, and other sources



From: Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. *BMJ* 2021;372:n71. doi: 10.1136/bmj.n71. For more information, visit: <http://www.prisma-statement.org/>

7.7 Appendix G Periodontal health results

No papers examined periodontal disease

7.8 Appendix H Complete quality assessment scores

Table 37 National Heart, Lung, and Blood Institute’s (NHLBI’s) quality assessment scores for observational cohort and cross-sectional studies for Question 2A

| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6 | 7. | 8 | 9 | 10 | 11 | 12 | 13 | 12 | 13 | 14 |
|---------------------|------|-------------|------------------------|-----|-----|------------------|-----|----------------|------------------|------------------|-----|------------------|----------------|----------------|----------------|----------------|------------------|----------------|-----------|
| Williams and Zwemer | 1990 | USA | Cross-sectional survey | Yes | Yes | No | Yes | Not applicable | Cannot determine | Cannot determine | Yes | Yes | Yes | Not applicable | Yes | Not applicable | Cannot determine | Not applicable | Partial |
| Riordan | 1993 | Australia | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | No | Yes | Yes | Not applicable | Cannot determine | Not applicable | Partial |
| Riordan | 2002 | Australia | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | No | Yes | Yes | Not applicable | Cannot determine | Not applicable | Partial |
| Clark et al. | 1994 | Canada | Cross-sectional survey | No | Yes | No | Yes | No | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | Not applicable | Not applicable | Not reported | Not applicable | Partial |
| Clark et al. | 1995 | Canada | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Cannot determine | Yes | Cannot determine | Yes | Not applicable | Not applicable | Not applicable | Not reported | Not applicable | Partial |
| Clark et al. | 2006 | Canada | Cross-sectional survey | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | Not applicable | Not applicable | Not reported | Not applicable | Extensive |
| Rock and Sabieha | 1997 | England, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | Cannot determine | Not applicable | Some |
| Kumar and Swango | 1999 | USA | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | Cannot determine | Not applicable | Partial |
| Tabari et al. | 2000 | England, UK | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | Yes | Not applicable | Yes | Not applicable | Some |
| Tiano et al. | 2009 | Brazil | Cross-sectional survey | Yes | Yes | Yes | Yes | No | Cannot determine | Cannot determine | Yes | Cannot determine | Yes | Not applicable | Not applicable | Not applicable | Not reported | Not applicable | None |
| de Moura et al. | 2013 | Brazil | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | No | Yes | Yes | Yes | Yes | No | Not applicable | Yes | Not applicable | Yes | Not applicable | Partial |
| Silva et al | 2021 | Brazil | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | No | Yes | Not applicable | Not applicable | Not reported | Not applicable | Partial |

HRB Document Template

| | | | | | | | | | | | | | | | | | | | |
|-----------------|------|-----------|------------------------|-----|-----|------------------|-----|-----|-----|-----|-----|-----|------------------|----------------|-----|----------------|------------------|----------------|---------|
| Bal et al. | 2015 | Australia | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | No | Not applicable | Yes | Not applicable | Cannot determine | Not applicable | Some |
| James et al. | 2021 | Ireland | Cross-sectional survey | Yes | Yes | Cannot determine | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | No | Not applicable | Partial |
| Marques et al. | 2021 | Brazil | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | Cannot determine | Not applicable | Partial |
| McLaren et al. | 2021 | Canada | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Not applicable | Not applicable | Not applicable | Partial |
| Mohd Nor et al. | 2021 | Malaysia | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | Not applicable | Not applicable | Not applicable | Partial |

*1. Research question stated

2. Study population clearly specified

3. Participation rate at least 50%

4. Subjects selected from the same population and inclusion and exclusion criteria prespecified

5. Sample size justification, power description, or variance and effect estimates provided

6. Exposure(s) of interest measured prior to outcome(s) measure

7. Timeframe sufficient to see an association between exposure and outcome

8. For exposures, study examine different levels of the exposure as related to the outcome

9. Exposure measures defined, valid, reliable, and consistently applied

10. Exposure(s) assessed more than once

11. Outcome measures defined, valid, reliable, and consistently applied: Caries

11. Outcome measures defined, valid, reliable, and consistently applied: Fluorosis

12. Outcome assessors blinded to the exposure status

13. Loss to follow-up 20% or less

14. Potential confounding exposures measured and adjusted statistically in outcomes

Table 38 National Heart, Lung, and Blood Institute's (NHLBI's) quality assessment scores for observational case-control studies for Question 2A

| Study ID | Year | Country | 1. Research question appropriate | 2. Study population specified | 3. Cases represent the cases in target population? | 4. Sample size justification? | 5. Controls selected from the same population | 6. Definitions, inclusion and exclusion criteria for select cases and controls valid, reliable, and implemented consistently | 7. Cases differentiated from controls | 8. Cases and/or controls randomly selected or census | 9. Use of concurrent controls | 10. Confirm that the exposure/risk occurred prior to the condition or event | 11. Measures of exposure clearly defined, valid, reliable, and implemented consistently | 12. Assessors of exposure blinded | 13. Potential confounding variables measured and adjusted statistically |
|------------------------------|------|---------|----------------------------------|-------------------------------|--|-------------------------------|---|--|---------------------------------------|--|-------------------------------|---|---|-----------------------------------|---|
| Osujp et al. | 1988 | Canada | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Yes | Partial |
| Keller Celeste and Blaya Luz | 2016 | Brazil | Yes | Yes | Yes | No | Yes | Yes | Yes | Yes | Yes | No | Yes | No | Some |

For each paper reporting on a longitudinal cohort study, cross-sectional survey, or case-control study, the scores were summed (for a total score ranging from 0.0 to 5.0). Papers scoring less than 3.0 were rated 'low quality', papers scoring 3.0 were rated 'moderate quality', and papers scoring 3.5 or more were rated 'high quality'. As many studies were cross-sectional in nature (point-in-time surveys) and scored 0.0 on item 13 (loss to follow-up not applicable), the maximum possible score for papers reporting on these types of studies was effectively capped at 4.0; for this reason, the threshold for 'high quality' was set at 3.5, rather than 4.0, in order to allow more effective differentiation of papers at the upper end of the range of scores. We also report the quality deficiencies by low-, moderate- and high-quality papers.

7.9 Appendix I Feasibility assessment results

Table 39 Feasibility assessment to determine the validity of meta-analysis to determine the summary effect of exposure to CWF and fluoridated toothpaste during the first 6 years of life on the prevalence of mild to severe dental fluorosis

| Study, year, country | Study design | Study population | CWF ppm (intervention) | Details of comparator | Outcome, measure, and cut offs | Statistical measure | Variance | Adjustment for confounding | Confounders | Study quality | Suitable for meta-analysis |
|-----------------------------------|-------------------------------|--|---|--|---|---|----------|---|---|---------------|--|
| Williams and Zwemer 1990 USA | Cross-sectional survey/census | 12–14-year-old school children | CWF 0.9 to 1.2 ppm plus preschool fluoridated toothpaste | 0.2 to 0.9 ppm plus fluoridated toothpaste | Dental fluorosis Tooth Surface Index of Fluorosis Single scores for each grade of fluorosis | Number and proportion reported for each score | No | Simple chi square tests and none were significant | Gender, race, preschool dietary patterns, preschool fluoridated toothpaste, swallowing toothpaste, and fluoride supplements | Low | No, due to type of comparator. In addition, the analysis does not investigate interactions between or control for confounding effects |
| Riordan 1993 Australia | Cross-sectional survey | Children born in 1983 (7 years of age) | Years exposed to CWF 0.8 ppm plus fluoridated toothpaste | Years exposed to CWF | Dental fluorosis Thylstrup and Fejerskov Index Scores 0–3 | Adjusted OR and 95% CI | Yes | Logistic regression | Age of weaning, preschool fluoridated toothpaste, swallowing toothpaste, and supplements | Low | No, as the comparator is not comparable with other comparators. In addition, the analysis does not investigate interactions between or control for confounding effects |
| Rock and Sabieha 1997 England, UK | Cross-sectional survey | School children aged 8–9-year-old | CWF 1.0 ppm plus low, medium and high fluoride toothpaste | Comparisons within toothpaste | Dental fluorosis Modified Thylstrup and Fejerskov Index Scores 0–6 0 versus 1–6 | Number and proportion reported for each score means and ranges for variables) by fluorosis status | Range | Interaction of paste weight and the type of brush, and brushing frequency and the occurrence of fluorosis | Regular or low fluoridated toothpaste and toothbrushing practices (age, toothpaste weight, times brushed teeth daily, fluoride in grammes ingested daily), and DMFT | Low | No, as the comparator is not comparable with other comparators. No adjusted statistical measures as interaction between significant tooth paste and brush variables |

| | | | | | | | | | | | |
|---------------------------------------|------------------------|--|--|--|---|--------------------------|--------|--|---|----------|---|
| Kumar and Swango 1999 USA | Cross-sectional survey | School children with lifelong residency, aged 7–10-years and 11–14-years | CWF 1±0.2 ppm plus fluoridated toothpaste | Nonfluorinated water plus fluoride toothpaste | Dental fluorosis Dean's Index of Fluorosis Normal compared with questionable, very mild, and mild to severe | Adjusted mean DMFS (SD) | SD | Regression models to determine factors associated with fluorosis | Age, race, poverty level, education level of household head, fluoridated toothpaste use before 2 years of age, supplements during first 8 years, and sealants | Low | No as interactions between 3–4 variables in models |
| Tabari <i>et al.</i> 2000 England, UK | Cross-sectional survey | 8–9-year-old school children who were lifetime residents in the area | CWF 1.0 ppm plus fluoride toothpaste | Less than 0.1 ppm plus fluoride toothpaste | Dental fluorosis Thylstrup and Fejerskov Index 0 compared with 1, 2, and 3+ | OR and 95% CI | 95% CI | Logistic regression model analysing the contribution of three variables to fluorosis— the area of residence (p <0.001), Jarman (deprivation) score (p =0.03), and type of toothpaste used (p =0.02) No interactions identified | Age started to brush Brushing frequency Amount of paste Toothpaste weight Toothpaste type Jarman score | Moderate | Yes |
| de Moura <i>et al.</i> 2013 Brazil | Cross-sectional survey | 8–12-year-old children who were lifelong residents of Teresina | CWF (0.6–0.8 ppm) status ascertained plus fluoridated toothpaste and tooth brushing educational programme | CWF 0.6–0.8 ppm plus fluoridated toothpaste | Dental fluorosis Thylstrup and Fejerskov Index 0 versus 1–6 | Adjusted ORs and 95% CI | 95% CI | Logistic regression to determine what prevents fluorosis | Age matched Adjusted for education and gender Group 1: Toothpaste (F content of 1,000 ppm) and taught toothbrushing practices | Low | No, not measuring CWF and fluoride toothpaste but the effect of education on prevention of fluorosis |
| Bal <i>et al.</i> 2015 Australia | Cross-sectional survey | School children aged 7–11 years | CWF 1 ppm 64% had lifetime exposure to CWF Fluoridated toothpastes, supplements, mouth rinses, and fluoride gel; Tooth brushing practices and age at exposure to | Control region (fluoridated at 1 ppm since 1967/9) | Dental fluorosis Dean's index of fluorosis Normal and questionable compared very mild to severe | Adjusted ORs and 95% CIs | 95% CI | Logistic regression to determine what prevents fluorosis | Five of 58 variables were significant: frequency of toothbrushing, rinsing habit after brushing, eating or licking toothpaste (these behaviours relate to when toothbrushing commenced as a | Low | No, as interactions present. Exposure to fluoridated water and water from various sources used for reconstitution of infant formula were highly |

| | | | | | | | | | | | |
|----------------------------------|----------------------------|---|---|--|---|--------------------------|----------------|---|---|------|---|
| | | | each fluoride intervention | | | | | | habit), exposure to fluoridated water, and type of water used for the reconstitution of infant formula Exposure to fluoridated water and water from various sources used for reconstitution of infant formula were highly correlated variables | | correlated variables |
| Celeste and Luz 2016 Brazil | Matched case-control study | 12-year-old schoolchildren | CWF 0.6–0.8 ppm Tooth brushing practices and age at exposure to each fluoride intervention. Swallowing toothpaste | Not applicable Cases had fluorosis and controls had no fluorosis | Dental fluorosis Dean's index of fluorosis Fluorosis cases versus no fluorosis controls | Adjusted ORs and 95% CIs | 95% CIs | Matched by sex and school case control study Conditional logistic regression to identify exposures in exposure cases Significant interaction between eating toothpaste, amount applied to brush, and size of toothbrush | Toothpastes, toothbrushing habits, supplements, mouth rinses, and fluoride gel. | Low | No as different study design and low quality |
| James <i>et al.</i> 2021 Ireland | Cross-sectional survey | Random sample of 5-year-old schoolchildren in Dublin & Cork-Kerry in 2014, follow up at age 8 years in 2017 | Reduced fluoride in CWF from 0.8–1 ppm in 2007 to 0.6–0.8 ppm since; plus fluoridated toothpaste and guidance | 0.2 ppm in rural Cork-Kerry with fluoride deficient water ; plus fluoridated toothpaste and guidance | Dental fluorosis Thylstrup and Fejerskov Index Normal and questionable versus very mild or higher | Not applicable | Not applicable | No significant findings for fluorosis outcome | National tooth brushing guidance in Urban Cork, Kerry and Dublin (since 2002) | High | No as not all data available No significant findings but no table informing us what was tested |

| | | | | | | | | | | | |
|--|-------------------------------|--|--|--|--|--------------------------------|---------|--|--|------|---|
| Marques <i>et al.</i> 2021 Brazil | Cross- sectional survey | High school students from 17 to 20 years of age, enrolled in public schools | Fluoridated toothpaste assumed plus CWF 0.6–0.8 ppm, | Fluoridated toothpaste assumed and fluoride deficient areas of Teresina | Dental fluorosis Thylstrup and Fejerskov Index Score 0 (absent) compared with scores 1 and 2 (very mild), scores 3 and 4 (moderate), and scores 5–9 (severe) | Adjusted ORs and 95% CIs | 95% CIs | In the final multivariate model, students exposed to fluoridated water were more likely to have very mild/mild fluorosis (OR 2.26; 95% CI 1.54–3.32) and moderate fluorosis (OR 3.66; 95% CI 1.93–6.95), than those who were not exposed. The odds of moderate fluorosis were 2.01 times higher in males than females | Model adjusted toothache, treatment need, how long since last appointment (years) and last appointment reason. Tooth brushing practices asked about. | High | No as measuring the additional influence of CWF over toothpaste on diagnosis of fluorosis, rather than the other way round |
|--|-------------------------------|--|--|--|--|--------------------------------|---------|--|--|------|---|

8 Appendices Question 2B

8.1 Appendix A Search Question 2B

Please see 7.1 for the search strategies and resources used in retrieving evidence for 2B. The same search results were used for both questions and a different screening code was utilised in screening the evidence. See Table 28 for a summary of search results for Q2B.

8.2 Appendix B PRISMA checklist and PRISMA-S for Question 2B

8.2.1 PRISMA checklist for Question 2B

| Topic | Item | Checklist item | Location where item is reported |
|-------------------------|------|--|---|
| TITLE | | | |
| Title | 1 | Identify the report as a systematic review. | Title page |
| ABSTRACT | | | |
| Abstract | 2 | See the PRISMA 2020 for Abstracts checklist. | Executive summary |
| INTRODUCTION | | | |
| Rationale | 3 | Describe the rationale for the review in the context of existing knowledge. | Section 1.1.4 |
| Objectives | 4 | Provide an explicit statement of the objective(s) or question(s) the review addresses. | Section 1.2 |
| METHODS | | | |
| Eligibility criteria | 5 | Specify the inclusion and exclusion criteria for the review and how studies were grouped for the syntheses. | Sections 2.3.1, 2.3.2, and 2.3.3 |
| Information sources | 6 | Specify all databases, registers, websites, organisations, reference lists and other sources searched or consulted to identify studies. Specify the date when each source was last searched or consulted. | Sections 2.4 and 2.5 |
| Search strategy | 7 | Present the full search strategies for all databases, registers and websites, including any filters and limits used. | Appendix A of Section 8, Sections 2.4 and 2.5 |
| Selection process | 8 | Specify the methods used to decide whether a study met the inclusion criteria of the review, including how many reviewers screened each record and each report retrieved, whether they worked independently, and if applicable, details of automation tools used in the process. | Section 2.6 Error! Reference source not found. |
| Data collection process | 9 | Specify the methods used to collect data from reports, including how many reviewers collected data from each report, whether they worked independently, any processes for obtaining or confirming data from study investigators, and if applicable, details of automation tools used in the process. | Section 2.7 |

| Topic | Item | Checklist item | Location where item is reported |
|-------------------------------|------|---|---------------------------------|
| Data items | 10a | List and define all outcomes for which data were sought. Specify whether all results that were compatible with each outcome domain in each study were sought (e.g. for all measures, time points, analyses), and if not, the methods used to decide which results to collect. | Sections 2.3.2, Table 3 |
| | 10b | List and define all other variables for which data were sought (e.g. participant and intervention characteristics, funding sources). Describe any assumptions made about any missing or unclear information. | Sections 2.7.1 |
| Study risk of bias assessment | 11 | Specify the methods used to assess risk of bias in the included studies, including details of the tool(s) used, how many reviewers assessed each study and whether they worked independently, and if applicable, details of automation tools used in the process. | Section 2.8 |
| Effect measures | 12 | Specify for each outcome the effect measure(s) (e.g. risk ratio, mean difference) used in the synthesis or presentation of results. | Section 2.9.1 |
| Synthesis methods | 13a | Describe the processes used to decide which studies were eligible for each synthesis (e.g. tabulating the study intervention characteristics and comparing against the planned groups for each synthesis (item #5)). | Section 2.9.2 |
| | 13b | Describe any methods required to prepare the data for presentation or synthesis, such as handling of missing summary statistics, or data conversions. | Section 2.9.3 |
| | 13c | Describe any methods used to tabulate or visually display results of individual studies and syntheses. | Section 2.9.2 and 2.9.3 |
| | 13d | Describe any methods used to synthesize results and provide a rationale for the choice(s). If meta-analysis was performed, describe the model(s), method(s) to identify the presence and extent of statistical heterogeneity, and software package(s) used. | Section 2.9.3 |
| | 13e | Describe any methods used to explore possible causes of heterogeneity among study results (e.g. subgroup analysis, meta-regression). | Section 2.9.3 |
| | 13f | Describe any sensitivity analyses conducted to assess robustness of the synthesized results. | Section 2.9.3 |
| Reporting bias assessment | 14 | Describe any methods used to assess risk of bias due to missing results in a synthesis (arising from reporting biases). | Section 2.9.2 and 2.9.3 |

| Topic | Item | Checklist item | Location where item is reported |
|-------------------------------|------|--|--|
| Certainty assessment | 15 | Describe any methods used to assess certainty (or confidence) in the body of evidence for an outcome. | Section 2.10 |
| RESULTS | | | |
| Study selection | 16a | Describe the results of the search and selection process, from the number of records identified in the search to the number of studies included in the review, ideally using a flow diagram. | Section 3.3, Appendix F of Section 8.6 |
| | 16b | Cite studies that might appear to meet the inclusion criteria, but which were excluded, and explain why they were excluded. | Appendix C of Section |
| Study characteristics | 17 | Cite each included study and present its characteristics. | Section 3.3.2, Table 48 |
| Risk of bias in studies | 18 | Present assessments of risk of bias [and/or quality assessment] for each included study. | Section 3.3.3, Tables 49 and 50, Appendix H of Section 8 |
| Results of individual studies | 19 | For all outcomes, present, for each study: (a) summary statistics for each group (where appropriate) and (b) an effect estimate and its precision (e.g. confidence/credible interval), ideally using structured tables or plots. | Section 3.3.4 |
| Results of syntheses | 20a | For each synthesis, briefly summarise the characteristics and risk of bias among contributing studies. | Section 3.3.3, Appendix H of Section 8 |
| | 20b | Present results of all statistical syntheses conducted. If meta-analysis was done, present for each the summary estimate and its precision (e.g. confidence/credible interval) and measures of statistical heterogeneity. If comparing groups, describe the direction of the effect. | Section 3.3.4.1.4, 3.3.4.1.5, 3.3.4.2.3 and 3.3.4.2.4 |
| | 20c | Present results of all investigations of possible causes of heterogeneity among study results. | Sections 3.3.4.1.4 and 3.3.4.2.4 |
| | 20d | Present results of all sensitivity analyses conducted to assess the robustness of the synthesized results. | Not applicable |
| Reporting biases | 21 | Present assessments of risk of bias due to missing results (arising from reporting biases) for each synthesis assessed. | Not applicable as mainly cross section surveys |

| Topic | Item | Checklist item | Location where item is reported |
|--|------|--|---|
| Certainty of evidence | 22 | Present assessments of certainty (or confidence) in the body of evidence for each outcome assessed. | Section 3.3.4.1.6 and 3.3.4.2.5 |
| DISCUSSION | | | |
| Discussion | 23a | Provide a general interpretation of the results in the context of other evidence. | Sections 4.1 and 4.2 |
| | 23b | Discuss any limitations of the evidence included in the review. | Section 4.3 |
| | 23c | Discuss any limitations of the review processes used. | Section 4.3 |
| | 23d | Discuss implications of the results for practice, policy, and future research. | Sections 4.4 and 4.5 |
| OTHER INFORMATION | | | |
| Registration and protocol | 24a | Provide registration information for the review, including register name and registration number, or state that the review was not registered. | Section 2.3.2 |
| | 24b | Indicate where the review protocol can be accessed, or state that a protocol was not prepared. | Section 2.3.2 |
| | 24c | Describe and explain any amendments to information provided at registration or in the protocol. | Section 2.3.2 |
| Support | 25 | Describe sources of financial or non-financial support for the review, and the role of the funders or sponsors in the review. | Not applicable as all authors are salaried public servants who are funded from the DOH public funding and are obliged to be objective |
| Competing interests | 26 | Declare any competing interests of review authors. | None |
| Availability of data, code and other materials | 27 | Report which of the following are publicly available and where they can be found: template data collection forms; data extracted from included studies; data used for all analyses; analytic code; any other materials used in the review. | Appendix D and E of Section 8 |

Source: Page *et al.* (2021)[1]

8.2.2 PRISMA-S Q 2 A and B

| Section/topic | # | Checklist item | Location(s) Reported |
|--|----|--|--|
| INFORMATION SOURCES AND METHODS | | | |
| Database name | 1 | Name each individual database searched, stating the platform for each. | Sections 2.4.4 and 2.5 |
| Multi-database searching | 2 | If databases were searched simultaneously on a single platform, state the name of the platform, listing all of the databases searched. | n/a |
| Study registries | 3 | List any study registries searched. | Sections 2.4 and 2.5 |
| Online resources and browsing | 4 | Describe any online or print source purposefully searched or browsed (e.g., tables of contents, print conference proceedings, web sites), and how this was done. | Sections 2.4.5, 2.4.6 |
| Citation searching | 5 | Indicate whether cited references or citing references were examined, and describe any methods used for locating cited/citing references (e.g., browsing reference lists, using a citation index, setting up email alerts for references citing included studies). | Section 2.4.5 |
| Contacts | 6 | Indicate whether additional studies or data were sought by contacting authors, experts, manufacturers, or others. | n/a |
| Other methods | 7 | Describe any additional information sources or search methods used. | n/a |
| SEARCH STRATEGIES | | | |
| Full search strategies | 8 | Include the search strategies for each database and information source, copied and pasted exactly as run. | Appendix A of Section 7 |
| Limits and restrictions | 9 | Specify that no limits were used, or describe any limits or restrictions applied to a search (e.g., date or time period, language, study design) and provide justification for their use. | Sections 2.5 and Appendix A of Section 7 |
| Search filters | 10 | Indicate whether published search filters were used (as originally designed or modified), and if so, cite the filter(s) used. | n/a |
| Prior work | 11 | Indicate when search strategies from other literature reviews were adapted or reused for a substantive part or all of the search, citing the previous review(s). | n/a |

| | | | |
|-------------------------|--------|--|-------------------------|
| Updates | 1 2 | Report the methods used to update the search(es) (e.g., rerunning searches, email alerts). | Appendix A of Section 7 |
| Dates of searches | 1 3 | For each search strategy, provide the date when the last search occurred. | Appendix A of Section 7 |
| PEER REVIEW | | | |
| Peer review | 1 4 | Describe any search peer review process. | Section 2.4.3 |
| MANAGING RECORDS | | | |
| Total Records | 1 5 | Document the total number of records identified from each database and other information sources. | Appendix A of Section 7 |
| Deduplication | 1 6 | Describe the processes and any software used to deduplicate records from multiple database searches and other information sources. | Section 2.4.4 |

PRISMA-S: An Extension to the PRISMA Statement for Reporting Literature Searches in Systematic Reviews

Rethlefsen ML, Kirtley S, Waffenschmidt S, Ayala AP, Moher D, Page MJ, Koffel JB, PRISMA-S Group.

Last updated February 27, 2020.

8.3 Appendix C Studies excluded at full text and extraction screening stages

8.3.1 Exclude on population

Exclude on population (n=19)

Bagramian RA. A 5-year school-based comprehensive preventive program in Michigan, U.S.A.

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8.4 Appendix D Extraction form

Table 40 General information data form

| Study ID | Author | Year | Location | Area | Objective | Secondary publication | Associated papers | Study design | Participant age | Artificial fluoridation | Fluoride interventions | Outcome | Outcome details | Extracted | Validated |
|-----------|--------------|----------------|----------|------------------------|--------------|--|---|--------------|-----------------------------------|------------------------------------|------------------------|------------------------------|---------------------------------|-----------|-----------|
| From Eppi | First author | Of publication | Country | State/County/City/Town | Aim of study | Data will not be extracted unless additional endpoints | Same overall project different analysis | HRB decision | Mean or ranges described in study | Confirm if explicitly stated (Y/N) | | Oral health outcome assessed | Including method of measurement | | |
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Table 41 Study design data form

| Study ID | Author | Year | Study design (Author allocated) | Study design (HRB allocated) | Justification | Length of study | Length of exposure to CWF | Details of exposure | Details of comparator | Eligibility criteria | Sample size calculation | Response rate | Blinding of assessors to exposure | % Lost to follow-up | Method for handling missing data | Data collection | Confounders | Control for confounding | Identification of effect modification | Effect modifiers | Notes |
|-----------|--------------|----------------|---------------------------------|------------------------------|---------------|-----------------|--|---------------------|-----------------------|----------------------|--|---------------|-----------------------------------|---------------------|---------------------------------------|-------------------|-------------|-------------------------|---------------------------------------|------------------|-------|
| From Eppi | First author | Of publication | As stated in the study | As agreed by research team | | | Length of time exposed to community water fluoridation | Including dose | Including dose | | expected prevalence, power to detect a difference and allowed variance, results CIs calculated | | | For main analysis | e.g. last observation carried forward | Brief description | | | Yes or not reported | | |
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Table 42 Study participants data form

| Study ID | Author | Year | Group for characteristics | N | Mean age/Age range | % Female | N included in final analysis |
|-----------|--------------|----------------|---------------------------|----------|--------------------|----------|------------------------------|
| From Eppi | First author | Of publication | | Enrolled | | | |
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Table 43 Outcomes

| Of publication | Outcome of interest: Caries | % caries primary teeth | % caries free primary teeth | % caries permanent teeth | % caries free permanent teeth | dmft/deft | dmfs/defs | DMFT | DMFS | Method of caries identification | Clinical examination criteria | Outcome of interest: Fluorosis | Fluorosis (Dean's index) | Fluorosis (Thylstrup-Fejerskov index) | Tooth Surfaces Index of Fluorosis | Type of teeth examined for fluorosis | Hypo mineralisation by photographs |
|----------------|-----------------------------|------------------------|-----------------------------|--------------------------|-------------------------------|-----------|-----------|------|------|---------------------------------|-------------------------------|--------------------------------|--------------------------|---------------------------------------|-----------------------------------|--------------------------------------|------------------------------------|
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Table 44 Caries outcome data form using example of primary dentition dmft

| Country | Author | Year | Age in years | CWF ppm | Baseline dmft CWF | Baseline CWF SD | Baseline CWF Total | Final dmft CWF | Final dmft SD CWF | Final CWF Total | Fluoride deficient ppm | Baseline mean dmft No F | Baseline SD No F | Baseline CWF Total | Final dmft No F | Final SD No F | Final No F Total | Difference in % point or dmft |
|---------|--------|------|--------------|---------|-------------------|-----------------|--------------------|----------------|-------------------|-----------------|------------------------|-------------------------|------------------|--------------------|-----------------|---------------|------------------|-------------------------------|
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This table was repeated for dmfs, % with CDC, and % without CDC for primary dentition. The table was also repeated for DMFT, DFMS, % with CDC, and % without CDC for permanent dentition

Table 45 Fluorosis outcome data form

| Country | Author | Year | Age in years | CWF ppm | Baseline % fluorosis | Baseline 95% CI | Baseline CWF affected number | Baseline CWF Total | Final % fluorosis | Final 95% CI | Final CWF affected number | Final CWF Total | Baseline 5 fluorosis No F | Baseline 95% CI No F | Baseline affected number No F | Baseline CWF Total | Final % No F | Final 95%CI No F | Final affected number No F | Final No F Total | Difference in % point or dmft |
|---------|--------|------|--------------|---------|----------------------|-----------------|------------------------------|--------------------|-------------------|--------------|---------------------------|-----------------|---------------------------|----------------------|-------------------------------|--------------------|--------------|------------------|----------------------------|------------------|-------------------------------|
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8.5 Appendix E Quality assessment tools

See 6.5 in Section 6.5 National Heart, Lung, and Blood Institute's (NHLBI's) quality assessment tool for observational cohort studies and cross-sectional surveys

Table 46 Bias domains included in version 2 of the Cochrane risk-of-bias tool for randomized trials for Question 2B

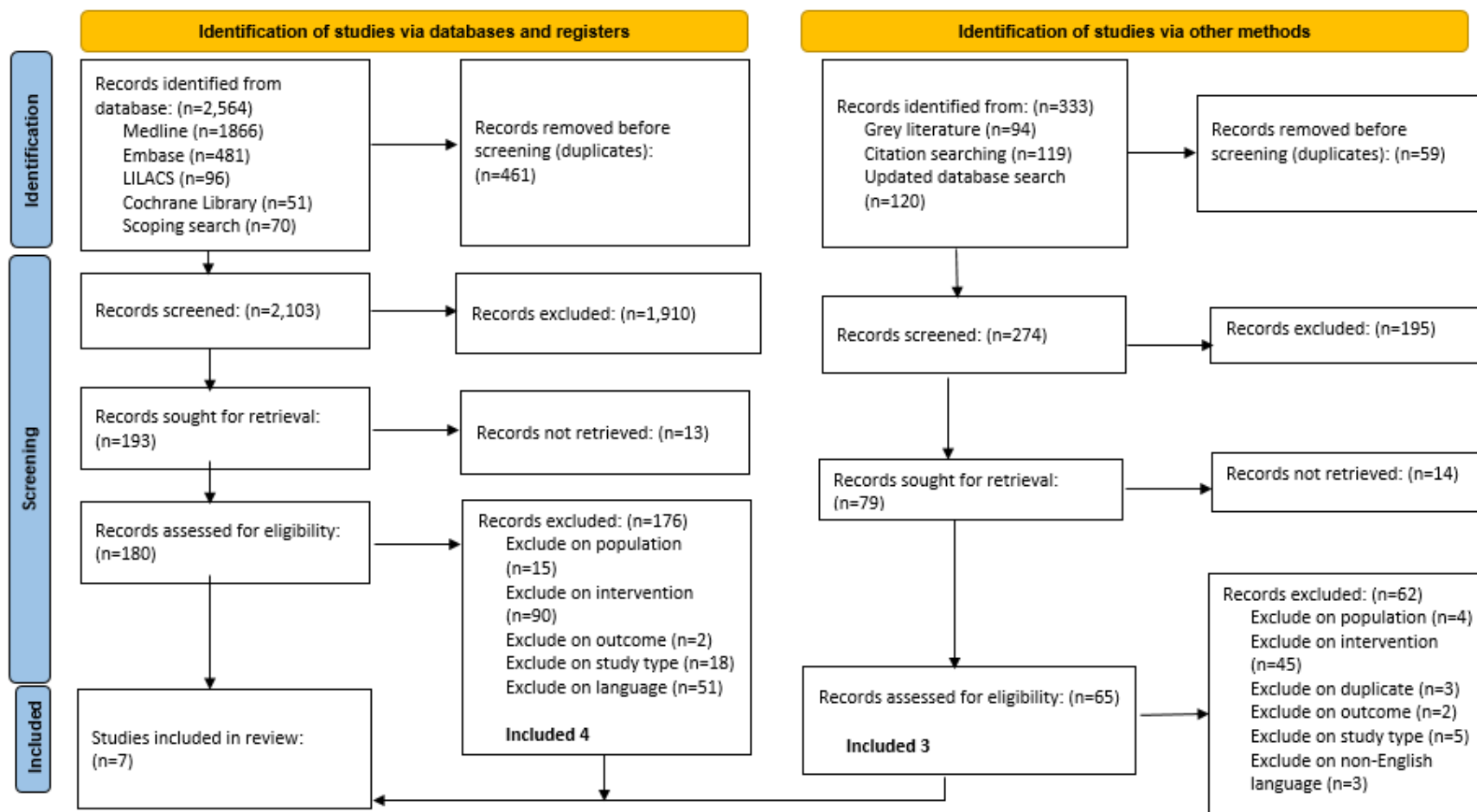
| Bias domain | Issues addressed* |
|--|---|
| Bias arising from the randomisation process | Whether: 1. The allocation sequence was random |
| | 2. The allocation sequence was adequately concealed, and 3. Baseline differences between intervention groups suggest a problem with the randomisation process. |
| Bias due to deviations from intended interventions | Whether: 4. Participants were aware of their assigned intervention during the trial 5. Carers and people delivering the interventions were aware of participants' assigned intervention during the trial 6. Deviations from the intended intervention arose because of the experimental context (i.e. do not reflect usual practice); and, if so, whether they were unbalanced between groups and likely to have affected the outcome |
| | 7. An appropriate analysis was used to estimate the effect of assignment to intervention, and, if not, whether there was potential for a substantial impact on the result 8. Important non-protocol interventions were balanced across intervention groups 9. Failures in implementing the intervention could have affected the outcome 10. Study participants adhered to the assigned intervention regimen, and 11. An appropriate analysis was used to estimate the effect of adhering to the intervention. |
| Bias due to missing outcome data | Whether: 12. Data for this outcome were available for all, or nearly all, participants randomised 13. Evidence that the result was not biased by missing outcome data, and 14. Missingness in the outcome was likely to depend on its true value (e.g. the proportions of missing outcome data, or reasons for missing outcome data, differ between intervention groups). |
| | Whether: 15. Method of measuring the outcome was inappropriate 16. Measurement or ascertainment of the outcome could have differed between intervention groups 17. Outcome assessors were aware of the intervention received by study participants, and |
| Bias in measurement of the outcome | |

| Bias domain | Issues addressed* |
|---|--|
| | 18. Assessment of the outcome was likely to have been influenced by knowledge of intervention received. |
| Bias in selection of the reported result | <p>Whether:</p> <p>19. Trial was analysed in accordance with a pre-specified plan that was finalised before unblinded outcome data were available for analysis</p> <p>20. Numerical result being assessed is likely to have been selected, on the basis of the results, from multiple outcome measurements within the outcome domain, and</p> <p>21. Numerical result being assessed is likely to have been selected, on the basis of the results, from multiple analyses of the data.</p> |

Adapted from [9]

8.6 Appendix F PRISMA flow diagram

PRISMA 2020 flow diagram for new systematic reviews which included searches of databases, registers, and other sources



From: Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. *BMJ* 2021;372:n71. doi: 10.1136/bmj.n71. For more information, visit: <http://www.prisma-statement.org/>

8.7 Appendix G Periodontal health results

No papers examined periodontal disease

8.8 Appendix H Complete quality assessment scores for Question 2B

Table 47 National Heart, Lung, and Blood Institute's (NHLBI's) quality assessment scores for observational cohort and cross-sectional studies for Question 2B

| Author | Year | Location | Study design | 1* | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 Caries | 11 Fluorosis | 12 | 13 | 14 |
|------------------|------|-----------|--|-----|-----|-----|-----|----------------|-----|-----|----------------|------------------|----------------|----------------|----------------|----------------|----------------|---------|
| Szpunar and Burt | 1988 | USA | Cross-sectional survey | Yes | Yes | No | Yes | No | Yes | Yes | Yes | Yes | Yes | Yes | Yes | No | Not applicable | Partial |
| Maupomé et al. | 2001 | Canada | Retrospective/prospective cohort study | Yes | Yes | No | Yes | No | Yes | Yes | Yes | Yes | Not reported | Yes | Not applicable | No | No | Partial |
| Clark et al. | 2006 | Canada | Cross-sectional survey | No | Yes | Yes | Yes | Not applicable | Yes | Yes | Yes | Cannot determine | Not applicable | Yes | Not applicable | Not applicable | Not applicable | Partial |
| McLaren et al. | 2021 | Canada | Cross-sectional survey | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Yes | Cannot determine | Yes | Yes | Not applicable | Not applicable | Not applicable | Partial |
| Bal et al. | 2015 | Australia | Cross-sectional survey | No | Yes | Yes | Yes | No | Yes | Yes | Not applicable | Yes | Not reported | Not applicable | Yes | Not reported | Not applicable | Partial |

*1. Research question stated

2. Study population clearly specified

3. Participation rate at least 50%

4. Subjects selected from the same population and inclusion and exclusion criteria prespecified

5. Sample size justification, power description, or variance and effect estimates provided

6. Exposure(s) of interest measured prior to outcome(s) measure

7. Timeframe sufficient to see an association between exposure and outcome

8. For exposures, study examine different levels of the exposure as related to the outcome

9. Exposure measures defined, valid, reliable, and consistently applied

10. Exposure(s) assessed more than once

11. Outcome measures defined, valid, reliable, and consistently applied: Caries

11. Outcome measures defined, valid, reliable, and consistently applied: Fluorosis

12. Outcome assessors blinded to the exposure status

13. Loss to follow-up 20% or less

14. Potential confounding exposures measured and adjusted statistically in outcomes

For each paper reporting on a longitudinal cohort study, cross-sectional survey, or case-control study, the scores were summed (for a total score ranging from 0.0 to 5.0). Papers scoring less than 3.0 were rated 'low quality', papers scoring 3.0 were rated 'moderate quality', and papers scoring 3.5 or more were rated 'high quality'. As many studies were cross-sectional in nature (point-in-time surveys) and scored 0.0 on item 13 (loss to follow-up not applicable), the maximum possible score for papers reporting on these types of studies was effectively capped at 4.0; for this reason, the threshold for 'high quality' was set at 3.5, rather than 4.0, in order to allow more effective differentiation of papers at the upper end of the range of scores. We also report the quality deficiencies by low-, moderate- and high-quality papers.

Table 48 RoB2 scores for randomised controlled trials included in Question 2B

| Country | Author | Year | Study design | Randomisation | Effect of assignment | Effect of adherence | Missing outcome data | Measurement of outcomes | Reported results | Overall ROB score |
|-----------|--------------|------|-----------------------------|---------------|----------------------|---------------------|----------------------|-------------------------|------------------|-------------------|
| Hong Kong | Jiang et al. | 2014 | Randomised controlled trial | Low | Some concerns | | Low | Low | Some concerns | Some concerns |
| Hong Kong | Lam et al. | 2021 | Randomised controlled trial | Low | Low | Low | Low | Some concerns | Low | Some concerns |

9 Appendices Question 3

9.1 Appendix A Q3 Search

Top level guidance from countries specified in the 087 protocol Australia, New Zealand, UK, Canada, Israel, USA, South American countries, EU countries. Community fluoridation countries only

Targeted searches for fluoridation guidance

Search date: 16 February 2022 and updated 9 February 2023

Browser: Firefox 97.0

Table 49 Organisational websites searched for Q3.

| Country | Website | URL |
|-----------|--|---|
| Australia | Australian Institute of Health and Welfare | https://www.aihw.gov.au/ |
| | Australian Dental Association | https://www.ada.org.au |
| | Government of Australia Department of Health | https://www.health.gov.au |
| | Australian Dental Association oral health information | https://www.teeth.org.au/ |
| | Australian Government National Health and Medical Research Council (NMHRC) | https://www.nhmrc.gov.au/ |
| | Australian Dental and Oral Health Therapists Association | https://www.adohta.net.au/ |
| | Safe Work Australia | https://www.safeworkaustralia.gov.au/ |
| | Royal Australian College of General Practitioners | https://www.racgp.org.au/ |
| | Health Direct (public health site) | https://www.healthdirect.gov.au/ |
| | Australian Research Centre for Population Oral Health | https://www.adelaide.edu.au/arcpoh/ |
| Brazil | Governo de Brasil | https://www.gov.br/ |
| | Governo de Brasil Ministério de Saúde | https://www.gov.br/saude/pt-br |
| | Ministério de Saúde. Secretaria de Atenção Primária à Saúde | https://aps.saude.gov.br |
| | Latin American Oral Health Association | https://laoha.org.br/ |
| | Pan American Health Organization Repository | https://iris.paho.org |
| Canada | Health Canada | https://www.canada.ca/en/health-canada.html |

| Country | Website | URL |
|----------------|---|--|
| Canada | Canadian Dental Association | https://www.cda-adc.ca/ |
| | CPG Infobase: Clinical Practice Guidelines | https://joulecma.ca/cpg/homepage |
| | Canadian Paediatric Society | https://cps.ca |
| | The Canadian Task Force on Preventive Health Care | https://canadiantaskforce.ca/ |
| | One Health Canada | https://www.onehealth.ca/ |
| | Canadian Association of Public Health Dentistry | https://caphd.ca/ |
| | Dental Hygiene Canada | https://www.dentalhygienecanada.ca/ https://www.dentalhygienecanada.ca/cdha/The_Profession_folder/Resources_folder/Position_Papers_Statements__Standards_folder/CDHA/The_Profession/Resources/Position_Statements.aspx |
| | Canadian Medical Association | https://www.cma.ca/ |
| | Canadian Foundation for Dental Hygiene Research and Education | https://www.cfdhre.ca/ |
| | Canadian Institute for Health Information | https://www.cihi.ca/en |
| Israel | State of Israel Department of Health | https://www.health.gov.il |
| | Israel Dental Association | https://www.ida.org.il/ |
| | State of Israel Government | https://www.gov.il/ |
| | Israeli Association of Pediatrics | https://pediatrics.doctoronly.co.il/ (full access not allowed) |
| | The Israel National Institute for Health Policy Research | http://www.israelhpr.org.il/en/ |
| New Zealand | New Zealand Ministry of Health | https://www.health.govt.nz/ |
| | New Zealand Dental Association | https://www.nzda.org.nz/ |
| | Office of the Prime Minister's Chief Science Officer | https://www.pmcsa.ac.nz |
| | Dental Council | https://www.dcnz.org.nz/ |
| | Dental Council Practice Standards | https://www.dcnz.org.nz/resources-and-publications/resources/practice-standards/ |
| United Kingdom | Gov.uk | https://www.gov.uk/ |

| Country | Website | URL |
|---------|---|---|
| | NICE | https://www.nice.org.uk/ |
| | NHS UK | https://www.nhs.uk |
| | British Dental Association | https://bda.org/ |
| | British Society of Paediatric Dentistry | https://www.bspd.co.uk/ |
| | British Society of Paediatric Dentistry Guidelines and Evidence Reviews | https://www.bspd.co.uk/Professionals/Resources/Clinical-Guidelines-and-Evidence-Reviews |
| | Royal College of Surgeons of England Faculty of Dental Surgery | https://www.rcseng.ac.uk/dental-faculties/fds |
| | British National Formulary | https://bnf.nice.org.uk |
| | College of General Dentistry (including Faculty of General Dental Practice) | https://cgdent.uk/ |
| | British Association for the Study of Community Dentistry | https://www.bascd.org/ |
| | Scottish Dental | https://www.scottishdental.org |
| | Scottish Government | https://www.gov.scot |
| | Public Health Scotland | https://publichealthscotland.scot/ |
| | Health Scotland [now Public Health Scotland] | http://www.healthscotland.com |
| | National Services Scotland Information Services Division [now Public Health Scotland] | https://www.isdscotland.org/ |
| | Public Health Information for Scotland | https://www.scotpho.org.uk/ |
| | ChildSmile | http://www.child-smile.org.uk/ |
| | Scottish Dental: information portal | https://www.scottishdental.org/ |
| | Scottish Intercollegiate Guideline Network | https://www.sign.ac.uk/ |
| | Scottish Dental Clinical Effectiveness programme | https://www.sdcep.org.uk/ |
| | SIGN | https://www.sign.ac.uk/ |
| | Welsh Government | https://gov.wales/ |
| | NHS Public Health Wales | https://phw.nhs.wales |
| | NHS 111 Wales | https://111.wales.nhs.uk/ |
| | Welsh Oral Health Information Unit | https://www.cardiff.ac.uk/research/explore/research-units/welsh-oral-health-information-unit |

| Country | Website | URL |
|------------------------------------|--|---|
| | Department of Health Northern Ireland | https://www.health-ni.gov.uk/ |
| | Department of Health Northern Ireland Professional dental guidance publications | https://www.health-ni.gov.uk/publications/professional-dental-guidance-publications |
| | HSC Public Health Agency | https://www.publichealth.hscni.net/ |
| | NI Direct | https://www.nidirect.gov.uk/ |
| | Health Service Executive for Northern Ireland | https://www.hseni.gov.uk/ |
| United States | American Dental Association | https://www.ada.org/ |
| | American Academy of Pediatric Dentistry (AAPD) | https://www.aapd.org/ |
| | My Children's Teeth (an information portal by the American Association of Pediatric Dentistry) | https://www.mychildrensteeth.org/ |
| | Centers for Disease Control and Prevention | https://www.cdc.gov/ |
| | American Public Health Association | https://apha.org/ |
| | US Preventive Service Task Force | https://www.uspreventiveservicestaskforce.org/uspstf/ |
| | Agency for Healthcare Research and Quality (AHRQ) | https://www.ahrq.gov/ |
| | U.S. Department of Health & Human Services (HHS) | https://www.hhs.gov/ |
| | American Academy of Pediatrics (AAP) | https://www.aap.org/ |
| | American Academy of Pediatrics: Oral health practice tools | https://www.aap.org/en/patient-care/oral-health/oral-health-practice-tools/ |
| | American Association of Public Health Dentistry | https://www.aaphd.org/ |
| | American Dietetics Association | https://www.eatright.org/ |
| General & international guidelines | Guidelines International Network (GIN) | https://guidelines.ebmportal.com/ |
| | Guideline Central | https://www.guidelinecentral.com/guidelines/ |
| | TRIP: Turning Research into Practice | https://www.tripdatabase.com/ |

| Country | Website | URL |
|------------------------------|--|---|
| | European Academy of Paediatric Dentistry (EAPD) | https://www.eapd.eu/ |
| | International Association of Paediatric Dentistry (IAPD) | https://iapdworld.org/ |
| | FDI World Dental Federation | https://www.fdiworldddental.org/ |
| | World Health Organization (WHO) | https://www.who.int/ |
| | Health Services/Technology Assessment Texts (HSTAT) | https://www.ncbi.nlm.nih.gov/books/NBK16710/ |
| Databases and search engines | General: EBSCO MEDLINE Complete | <p>PT "Guideline+") OR (MH "Guidelines as Topic+")) OR ((TI (guideline OR guidelines OR guidance OR standards OR "white paper" OR "policy statement" OR "best practice"))) OR ((MW (guideline OR guidelines OR guidance OR standards OR "white paper" OR "policy statement" OR "best practice")))</p> <p>AND</p> <p>(MH "Fluorides") OR (MH "Fluorides, Topical") OR ((TI (fluoride OR "oral health" OR caries OR tooth OR teeth OR "dental health" OR dentist* OR toothpaste OR dentifrice OR varnish* OR sealant*)) OR ((AB (fluoride OR "oral health" OR caries OR tooth OR teeth OR "dental health" OR dentist* OR toothpaste OR dentifrice OR varnish* OR sealant*))</p> <p>AND</p> <p>TI (boy# OR boyfrien* OR boyhood* OR child* OR fifth-grader* OR first-grader* OR fourth-grader* OR girl# OR girlfriend* OR girlhood* OR juvenil* OR kid# OR kindergarten* OR minor# OR minority OR paediatric* OR paediatric* OR pediatric* OR PICU OR preschool* OR pre-school* OR second-grader* OR seventh-grader* OR sixth-grader* OR stepchild* OR step-child* OR third-grader* OR toddler# OR young OR youngster* OR youth*) OR AB (boy# OR boyfrien* OR boyhood* OR child* OR fifth-grader* OR first-grader* OR fourth-grader* OR girl# OR girlfriend* OR girlhood* OR juvenil* OR kid# OR kindergarten* OR minor# OR minority OR paediatric* OR paediatric* OR pediatric* OR PICU OR preschool* OR pre-school* OR</p> |

| Country | Website | URL |
|---------|--|---|
| | | <p>second-grader* OR seventh-grader* OR sixth-grader* OR stepchild* OR step-child* OR third-grader* OR toddler# OR young OR youngster* OR youth*) OR MH ("Child" OR "Child, Preschool")</p> <p>AND</p> <p>limit to 2009-2023</p> <p>Fluoride/fluoridation/ toothpaste/varnish/dentifrice/sealant Dental/dentist/teeth/tooth Guidance/guide/guideline/summary/position statement/white paper</p> |
| | General: Google.com | |
| | General: Google Scholar | <p>Fluoride/fluoridation/ toothpaste/varnish/dentifrice/sealant Dental/dentist/teeth/tooth Guidance/guide/guideline/summary/position statement/white paper</p> |
| | Virtual Health Library (VHL) Regional Portal https://bvsalud.org/en/ | Fluor* |

10 GRADE scores and justifications

Table 50 GRADE scores and justifications for the primary dentition dental caries studies

| Outcome and number of studies* | Study design score | Study design justification | Participants numbers (High quality, moderate quality studies) | Risk of bias | Risk of bias justification | Inconsistency of results | Inconsistency of results justification | Indirectness | Indirectness justification | Imprecision | Imprecision justification | Publication bias | Publication bias justification | Final score† | Certainty of the evidence |
|---|--------------------|---|--|--------------|---|--------------------------|--|--------------|---|-------------|---|------------------|--|--------------|---------------------------|
| Primary dentition – dental caries | | | | | | | | | | | | | | | |
| dmft (single time point) - 18 studies (excludes 3 outliers) | -5 | Majority cross-sectional surveys (1 cohort study) | All 20782 High (H) 3521 Moderate (M) 13231 (64%) Low (L) 4030 (19%) | -2 | ≥ 75% participants in moderate or low quality studies | -2 | Heterogeneity = 97.1% | 0 | PICO framework was carefully designed to ensure relevance of included studies | 0 | Narrow CI, good sample size | 0 | Our search was comprehensive, findings are largely in favour of intervention | -9 | Very low |
| dmft (two time points) - 5 studies | -5 | Cross-sectional surveys | All 3225 H 912 M 1912 (59%) L 401 (12%) | -1 | < 75% participants in moderate or low quality studies | -2 | Heterogeneity based on single time point MA, inconsistent findings between studies | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | No CI available, some imprecision assumed | 0 | Our search was comprehensive, findings are largely in favour of intervention | -9 | Very low |
| dmfs (single time point) - 6 studies (excludes 1 outlier) | -5 | Cross-sectional surveys | All 3687 H 0 M 1933 (52%) L 1754 (48%) | -2 | ≥ 75% participants in moderate or low quality studies | -2 | Heterogeneity = 92.6% | 0 | PICO framework was carefully designed to ensure relevance of included studies | 0 | Narrow CI | 0 | Our search was comprehensive, findings are largely in favour of intervention | -9 | Very low |

| | | | | | | | | | | | | | | | |
|--|-----|-------------------------|---|-----|---|-----|--|-----|---|-----|---|-----|--|-----|----------|
| dmfs (two time points) - 0 studies | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A |
| % without CDC in primary dentition (single time point) - 4 studies | -5 | Cross-sectional survey | All 14582 H 912 642 (4%) 13028 (89%) | -2 | ≥ 75% participants in moderate or low quality studies | -2 | 84.0% | 0 | PICO framework was carefully designed to ensure relevance of included studies | 0 | Narrow CI | 0 | Our search was comprehensive, findings are largely in favour of intervention | -9 | Very low |
| % without CDC in primary dentition (two time points) - 1 study | -5 | Cross-sectional survey | All 3509 H 0 M 1166 (33%) L 2343 (67%) | -2 | ≥ 75% participants in moderate or low quality studies | 0 | Results based on a single study, reported by the authors to be statistically significant in favour of intervention | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | Wide CI at both time points | 0 | Our search was comprehensive, findings are largely in favour of intervention | -8 | Very low |
| % with CDC in primary dentition (single time point) - 4 studies | -5 | Cross-sectional surveys | All 3926 H 1717 M 2209 (56%) L 0 (0%) | -1 | < 75% participants in moderate quality studies | 0 | Heterogeneity = 0% | 0 | PICO framework was carefully designed to ensure relevance of included studies | 0 | Narrow CI, good sample size | 0 | Our search was comprehensive, findings are largely in favour of intervention | -6 | Low |
| % with CDC in primary dentition (two time points) - 2 studies | -5 | Cross-sectional surveys | All 1614 H 0 M 1614 (100%) L 0 (0%) | -2 | ≥ 75% participants in moderate quality studies | -2 | Percentage point differences were very different between studies | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | Narrow CI for each study, but small sample size | 0 | Our search was comprehensive, findings are largely in favour of intervention | -9 | Very low |

Table 51 GRADE scores and justifications for the permanent dentition dental caries studies

| Outcome and number of studies* | Study design score | Study design justification | Participants numbers (High quality, moderate quality studies) | Risk of bias | Risk of bias justification | Inconsistency of results | Inconsistency of results justification | Indirectness | Indirectness justification | Imprecision | Imprecision justification | Publication bias | Publication bias justification | Final score† | Certainty of the evidence |
|---|--------------------|----------------------------|---|--------------|---|--------------------------|--|--------------|---|-------------|--|------------------|--|--------------|---------------------------|
| Permanent dentition – dental caries | | | | | | | | | | | | | | | |
| DMFT (single time point) - 21 studies (excluded 4 outlier papers) | -5 | Cross-sectional surveys | All 17644 H 2065 M 10694 (61%) L 4885 (28%) | -2 | ≥ 75% participants in moderate or low quality studies | -2 | Heterogeneity = 98.4% | 0 | PICO framework was carefully designed to ensure relevance of included studies | 0 | Narrow CI, good sample size | 0 | Our search was comprehensive, findings are largely in favour of intervention | -9 | Very low |
| DMFT (two time points) - 5 studies | -5 | Cross-sectional surveys | All 4333 H 0 M 4333 (100%) L 0 (0%) | -2 | ≥ 75% participants in moderate quality studies | -2 | Heterogeneity based on single time point MA, inconsistent findings between studies | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | No CI available, some imprecision assumed | 0 | Our search was comprehensive, findings are largely in favour of intervention | -10 | Very low |
| DMFS (single time point) 5 studies (excluded 1 outlier paper) | -5 | Cross-sectional surveys | 19354 H 0 M 393 (2%) L 18961 (98%) | -2 | ≥ 75% participants in moderate or low quality studies | -2 | Heterogeneity = 98.5% | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | Narrow CIs for each study, but small sample size | 0 | Our search was comprehensive, findings are largely in favour of intervention | -10 | Very low |
| DMFS (two time points) - 0 studies | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A | N/A |

| Outcome and number of studies* | Study design score | Study design justification | Participants numbers (High quality, moderate quality studies) | Risk of bias | Risk of bias justification | Inconsistency of results | Inconsistency of results justification | Indirectness | Indirectness justification | Imprecision | Imprecision justification | Publication bias | Publication bias justification | Final score† | Certainty of the evidence |
|--|--------------------|----------------------------|---|--------------|---|--------------------------|--|--------------|---|-------------|----------------------------|------------------|--|--------------|---------------------------|
| % without CDC in permanent dentition (single time point) - 3 studies | -5 | Cross-sectional survey | All 9090 H 0 M 988 (11%) L 8102 (89%) | -2 | ≥ 75% participants in moderate or low quality studies | -2 | Heterogeneity = 96.6% | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | Wide CI | 0 | Our search was comprehensive, findings are largely in favour of intervention | -10 | Very low |
| % without CDC in permanent dentition (two time points) - 1 study | -5 | Cross-sectional survey | All 1079 H 0 M 1079 (100%) L 0 (0%) | -2 | ≥ 75% participants in moderate quality studies | 0 | Results based on a single study, reported by the authors to be statistically significant in favour of intervention | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | Wide CI, small sample size | 0 | Our search was comprehensive, findings are largely in favour of intervention | -8 | Very low |
| % with CDC in permanent dentition (single time point) - 3 studies | -5 | Cross-sectional surveys | All 3236 H 1703 M 1533 (47%) L 0 (0%) | -1 | <75% participants in moderate quality studies | -2 | Heterogeneity = 95% | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | Wide CI | 0 | Our search was comprehensive, findings are largely in favour of intervention | -9 | Very low |
| % with CDC in permanent dentition (two time points) - 2 studies | -5 | Cross-sectional surveys | All 1978 H 0 M 1978 (100%) L 0 (0%) | -2 | ≥ 75% participants in moderate quality studies | 0 | Two linked studies showed the percentage point difference strongly in favour of the intervention | 0 | PICO framework was carefully designed to ensure relevance of included studies | 0 | Narrow CIs for each study, | 0 | Our search was comprehensive, findings are largely in favour of intervention | -7 | Very low |

†There were no upgrades for large magnitude of effect, dose-gradient response or effect of plausible residual confounding

Table 52 GRADE scores and justifications for the dental fluorosis studies

| Outcome and number of studies | Study design score | Study design justification | Risk of bias | Risk of bias justification | Inconsistency of results | Inconsistency of results justification | Indirectness | Indirectness justification | Imprecision | Imprecision justification | Publication bias | Publication bias justification | Final score | Certainty of the evidence |
|---|--------------------|---|--------------|---|--------------------------|--|--------------|---|-------------|---------------------------|------------------|---|-------------|---------------------------|
| Q1 Fluorosis (26 studies in 33 papers) | -5 | 26 Cross-sectional surveys | -2 | ≥ 75% participants in moderate or low-quality studies | -2 | Different country contexts influence the baseline fluorosis prevalence | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | No CIs | 0 | Our search was comprehensive, findings are largely associated with the intervention | -10 | Very low |
| Q2A CWF plus fluoride toothpaste (17 papers) | -5 | 15 Cross-sectional survey, 2 case control studies | -2 | ≥ 75% participants in moderate or low-quality studies | -2 | Different country contexts influence the baseline fluorosis prevalence | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | Few CIs calculated | 0 | Our search was comprehensive, findings are largely associated with the intervention | -10 | Very low |
| Q2B CWF plus fluoride toothpaste plus other topical fluoride interventions (7 papers) | -5 | 4 cross-sectional surveys, 1 prospective cohort study and 2 block randomised trials | -2 | ≥ 75% participants in low quality studies | -2 | Different country contexts influence the baseline fluorosis prevalence | 0 | PICO framework was carefully designed to ensure relevance of included studies | -1 | Few CIs calculated | 0 | Our search was comprehensive, findings are largely associated with the intervention | -10 | Very low |

11 Comparison between Cochrane 2015 review and HRB 2024 review

Table 53 Fluorosis: Comparison of Cochrane 2015 and HRB included papers for Question 1

| Included by Cochrane | Found in HRB search | Reason for exclusion by HRB | Included by HRB for fluorosis |
|-----------------------------|---------------------|--|-------------------------------|
| Acharya 2005 | Yes | Natural endemic fluoride in India | No |
| Adair 1999 | Yes | Excluded at full text Mixed natural and CWF intervention Exclude | No |
| Al-Alousi 1975 | Yes | Excluded at extraction as assessing all enamel defects not just fluorosis | Yes |
| Alarcon-Herrera 2001 | Yes | Natural fluoride at high levels in Mexico | No |
| Albrecht 2004 | Yes | Natural fluoride at suboptimal, optimal and high levels in Hungary | No |
| AlDosari 2010 | Yes | Natural fluoridation of water to determine ideal level | No |
| Angelillo 1999 | Yes | Natural fluoridation of water | No |
| Arif 2013 | Yes | Natural fluoride at high levels in India | No |
| Azcurra 1995 | Yes | Natural fluoride at very low and very high levels in Argentina | No |
| Beltran-Aguilar 2002 | Yes | Natural, optimal and suboptimal in the USA. Optimal includes CWF, optimal, and adjusted fluoride. | No |
| Booth 1991 | Yes | Excluded for fluorosis as assessment of enamel defects in general. Included for dental caries | No |
| Brothwell 1999 | Yes | Varying levels of natural fluoride | No |
| Chandrashekar 2004 | Yes | Natural endemic fluoride in India | No |
| Chen 1989 | Yes | Drinking water contains negligible, optimal, and above-optimal concentrations of natural fluoride in China | No |
| Chen 1993 | Yes | Drinking water contains high concentrations of natural fluoride in China | No |

| | | | |
|-----------------------------|-----|---|-----|
| Clark 1993 | Yes | Included | Yes |
| Clarkson 1989 | Yes | Excluded as measurement of dental development effects (of which fluorosis is only 1) in Ireland and New Zealand | No |
| Cochran 2004a | Yes | Excluded on study design as review. Also, a mix of natural and CWF exposures in Europe | No |
| Correia Sampaio 1999 | Yes | Excluded as mix of natural and CWF fluoride in Brazil | No |
| Cutress 1985 | Yes | Excluded as defects of tooth enamel in New Zealand. | No |
| Driscoll 1983 | Yes | Excluded as optimal and above-optimal natural water fluoride concentrations in USA | No |
| Ekanayake 2002 | Yes | Excluded as natural water fluoride concentrations in Sri Lanka | No |
| Eklund 1987 | Yes | Excluded as optimal and above-optimal natural water fluoride concentrations in USA | No |
| Ellwood 1995 | Yes | Excluded for fluorosis as assessment of enamel defects in general. Included for dental caries | No |
| Ellwood 1996 | Yes | Excluded for fluorosis as assessment of enamel defects in general. Included for dental caries | No |
| Firempong 2013 | Yes | Excluded as high natural water fluoride concentrations in Ghana | No |
| Forrest 1965 | Yes | Excluded, CWF in intervention area in Wales, but ppm for control area not reported | No |
| Garcia-Perez 2013 | Yes | Excluded as natural fluoride plus salt fluoridation, not CWF in Mexico | No |
| Gaspar 1995 | No | Not located Gaspar M, Pereira A, Moreira B. Non-fluorosis and dental fluorosis opacities in areas with lower (0.2 ppm F) and optimum (0.7 ppm F) fluoride concentration in drinking water [Opacidades de esmalte de origem não fluorótica e fluorose dentária em áreas | No |

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|------------------------------|-----|---|-----|
| | | com baixa (0,2 ppm F) e ótima (0,7 ppm F) concentrações de flúor nas águas de abastecimento público]. Revista Brasileira de Odontologia 1995;52(2):13-8. We would exclude on foreign language | |
| Grimaldo 1995 | Yes | Excluded as natural fluoride in Mexico | No |
| Grobler 1986 | Yes | Excluded as optimal and high natural fluoride areas, ?South Africa | No |
| Grobler 2001 | Yes | Excluded as natural fluoride areas, ?South Africa | No |
| Haavikko 1974 | Yes | Natural fluoride | No |
| Heintze 1998 | Yes | Included CWF in 2 areas and fluoride deficient comparator | Yes |
| Heller 1997 | Yes | Excluded as intervention school water not home water. Both optimal natural and CWF included and combined | No |
| HernandezMontoya 2003 | Yes | Exclude as high natural fluoride in Mexico | No |
| Hong 1990 | Yes | Included | Yes |
| Ibrahim 1995 | Yes | Excluded as optimal and high natural fluoride areas in Sudan | No |
| Indermitte 2007 | Yes | Excluded as natural fluoride areas in Estonia | No |
| Indermitte 2009 | Yes | Excluded as natural fluoride (0.01 to 7.20 ppm) in Estonia | No |
| Ismail 1990 | Yes | Included | Yes |
| Jackson 1975a | Yes | Excluded at extraction as assessing all enamel defects not just fluorosis Included for dental caries | Yes |
| Jackson 1999 | Yes | Excluded as naturally fluoridated communities in Indiana, USA | No |
| Kanagaratnam 2009 | Yes | Excluded as no dose of fluoride for CWF. Diffuse opacity Diffuse opacities appear white when the tooth erupts, and have a similar range of translucency to the demarcated defects. The main difference is that they lack a margin | No |

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|---------------------------|-----|--|----------------------|
| | | which clearly defines them from the adjacent normal enamel. Fluoride induced lesions are usually found within this type (Suckling <i>et al.</i> , 1989). | |
| Kotecha 2012 | Yes | Exclude as high natural fluoride in India | No |
| Kumar 2007 | Yes | Exclude as high natural fluoride in India | No |
| Kunzel 1976 | Yes | Exclude as natural fluoride values of between 0.1 and 2.6 ppm in different rural areas of Cuba | No |
| Leverett 1986 | Yes | Excluded as optimal fluoride may mean adjusted natural fluoride, USA Query | No |
| Levine 1989 | Yes | Excluded on outcome enamel hypoplasia, Birmingham and Leeds and other causes besides fluorosis | No |
| Lin 1991 | No | Excluded natural endemic fluoride in China | No would not include |
| Louw 2002 | Yes | Excluded natural endemic fluoride, South Africa | No |
| Machiulskiene 2009 | Yes | Excluded natural endemic fluoride in Lithuania | No |
| Mackay 2005 | Yes | Excluded as no comparator ppm provided DDE diffuse opacity | No |
| Macpherson 2007 | Yes | Excluded natural endemic fluoride in Sweden | No |
| Mandinic 2009 | Yes | Excluded natural endemic fluoride in Serbia | No |
| Marya 2010 | Yes | Excluded natural endemic fluoride in India | No |
| Masztalerz 1990 | Yes | Excluded natural endemic fluoride in Poland | No |
| McGrady 2012 | Yes | Excluded as no comparator ppm provided | No |
| McInnes 1982 | Yes | Excluded natural endemic fluoride, South Africa | No |
| Mella 1992 | Yes | Excluded natural endemic fluoride in Chile | No |

| | | | |
|----------------------------|-----|---|-----------------------|
| Mella 1994 | Yes | Excluded natural endemic fluoride in Chile | No |
| Milsom 1990 | Yes | Excluded as enamel defects (diffuse separate), UK | No |
| Montero 2007 | Yes | Excluded natural endemic fluoride and salt in Venezuela | No |
| Nanda 1974 | Yes | Excluded natural endemic fluoride in India | No |
| Narbutaite 2007 | Yes | Excluded natural endemic fluoride in Lithuania | No |
| Narwaria 2013 | Yes | Excluded natural endemic fluoride in India | No |
| Nunn 1994a | Yes | Natural or CWF not clear Outcome DDE (diffuse) | No |
| Ockerse 1941 | Yes | Excluded natural endemic fluoride, South Africa | No |
| Pontigo-Loyola 2008 | Yes | Excluded as endemic fluoridated communities in Mexico | No |
| Ray 1982 | Yes | Excluded natural endemic fluoride in India | No |
| Riordan 1991 | Yes | Included | Yes |
| Riordan 2002 | Yes | Excluded as not examining the effects of CWF per se but low fluoride toothpaste and supplements. Q2 | No |
| Rwenyonyi 1998 | Yes | Excluded natural endemic fluoride in Uganda | No |
| Rwenyonyi 1999 | Yes | Excluded natural endemic fluoride in Uganda | No |
| Saravanan 2008 | Yes | Excluded natural endemic fluoride in India | No |
| Sellman 1957 | No | Excluded natural endemic fluoride in Sweden | No, would not include |
| Shanthi 2014 | Yes | Excluded natural endemic fluoride in India | No |
| Shekar 2012 | Yes | Excluded natural endemic fluoride in India | No |
| Stephen 2002 | Yes | Excluded naturally fluoridated and fluoride deficient townships of Scotland | No |

| | | | |
|-----------------------------|-----|--|-----------------------|
| Szpunar 1988 | Yes | Included Q 1and3 | Yes |
| Tabari 2000 | Yes | Included | Yes |
| Tsutsui 2000 | Yes | Excluded natural endemic fluoride in Japan | No |
| Wang 1993 | No | Excluded natural endemic fluoride in China | No, would not include |
| Wang 1999 | Yes | Excluded natural endemic fluoride in China | No |
| Wang 2012 | Yes | Excluded natural endemic fluoride in China | No |
| Warnakulasuriya 1992 | Yes | Excluded natural endemic fluoride in Sri Lanka | No |
| Warren 2001 | Yes | Excluded as Iowa cohort has a mix of natural fluoride and CWF, USA | No |
| Wenzel 1982 | Yes | Excluded natural endemic fluoride in Denmark | No |
| Wondwossen 2004 | Yes | Excluded natural endemic fluoride in Ethiopia | No |
| Zheng 1986 | Yes | Suitable but excluded on Language | No |
| Zimmermann 1954 | Yes | Excluded as natural fluoride in USA | No |

Table 54 Dental caries: Comparison of Cochrane 2015 and 2024 and HRB included papers for Question 1

| Included by Cochrane in 2015 and 2022 | Found in HRB search | Reason for exclusion by HRB | Included by HRB for dental caries |
|---------------------------------------|---------------------|---|-----------------------------------|
| Adriasola 1959 | Yes | Spanish language CWF intervention 1.0 ppm No ppm for control | No |
| Arnold 1956/1957 | Yes | Included | Yes |
| Ast 1951 | Yes | Included | Yes |
| Backer-Dirks 1961 | Yes | Included | Yes |
| Beal 1971 | Yes | Included | Yes |
| Beal 1981 | Yes | Included | Yes |
| Blinkhorn (unpublished)? 2015 | Yes | Neither intervention nor control levels of F reported | No |
| Brown 1965 | Yes | Included | Yes |
| DHSS England 1969 | Yes | Excluded as control ppm not stated | No |
| DHSS Scotland 1969 | Yes | Excluded as control ppm not stated | No |
| DHSS Wales 1969 | Yes | Excluded as control ppm not stated | No |
| Goodwin 2022 | Yes | Included | |
| Gray 2001 | Yes | Included | Yes |
| Guo 1984 | Yes | Included | Yes |
| Hardwick 1982 | Yes | Included | Yes |
| Kunzel 1997 | Yes | Included | Yes |
| Loh 1996 | Yes | HRB excluded as overview and included original study by Wong <i>et al.</i> | Yes |
| Pot 1974 | Yes | Excluded on foreign language (dutch) Have English publications for this cohort | No |
| Tessier 1987 | Yes | Excluded on foreign language (French) No control F ppm | No |
| Maupomé 2001 | Yes | Included | Yes |

