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Title: Investigating societal determinants of oral health – opportunities and challenges in multilevel studies

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Abstract:

The high prevalence of oral diseases and the persistent nature of socioeconomic inequalities in oral health outcomes across societies presents a significant challenge for public health globally. A debate exists in epidemiology on the merits of investigating population variations in health and its determinants over studying individual health and its individual risk factors. The choice of analytical unit for health outcomes at the population level has policy implications and consequences for the causal understanding of population-level variations in health/disease. There is a lack of discussion in oral epidemiology on the relevance of studying population variations in oral health. Evidence on the role of societal factors in shaping variations in oral health at both the individual-level, and the population-level, is also mounting. Multilevel studies are increasingly applied in social epidemiology to address hypotheses generated at different levels of social organization, but the opportunities offered by multilevel approaches are less applied for studying determinants of oral health at the societal level. Multilevel studies are complex as they aim to examine hypotheses generated at multiple levels of social organization and require attention to a range of theoretical and methodological aspects from the stage of design to analysis and interpretation. This discussion paper aims to highlight the value in studying population variations in oral health. It discusses the opportunities provided by multilevel approaches to study societal determinants of oral health. Finally, it reviews the key methodological aspects related to operationalizing multilevel studies of societal determinants of oral health.

1 Introduction

2 Oral diseases affect half of the global population.¹ Untreated dental caries remains as the
3 most prevalent non-communicable disease world-wide,¹ despite some success in reducing
4 childhood dental caries in high-income countries during the latter part of the twentieth
5 century.² Persistent socioeconomic inequalities in oral health outcomes within and between
6 societies are also highlighted.³ Collectively, the two issues reflect limited or inadequate
7 policy responses at both global and local levels. High levels of disease and inequalities in oral
8 health persist because current prevention methods that are based mainly on the biomedical
9 approach and focus on changing individual behaviour. These approaches tend to result in
10 only short term improvements for certain patients but do not address the underlying causes
11 of diseases.^{3, 4} Evidence-based actions at the population level are necessary to reduce the
12 enormous burden of oral diseases and counter the persistent socioeconomic inequalities in
13 oral health outcomes.

14 Majority of oral epidemiologic studies investigate only individual variations in oral health
15 and its individual determinants notwithstanding the current challenges in population oral
16 health.^{1, 3} Studies that examine individual-variations in oral health within populations and its
17 determinants tend to examine biologic/behavioral effects.⁵⁻⁷ The underlying biomedical and
18 individual-based approach of such epidemiological investigations is a key limitation as they
19 do not address the underlying causes of diseases. Individual-based approaches neglect
20 population variations in oral health and its societal risks. Alternatively, studies of population
21 variations in health are fundamental if the goal is to ascertain the determinants of population
22 health.⁵ These investigations are directed to study population-level variations in oral health
23 and its determinants, also called as ecologic effects.⁶ Studies of population variations in oral
24 health are also critical to understanding systematic differences due to which oral health
25 inequalities mirror the social inequalities present within societies. A timely discussion of the
26 theoretical rationale for investigating population variations in oral health is essential in the
27 light of current challenges.

28 Multilevel studies are progressively applied in the discipline of social epidemiology and in
29 oral health.⁸ The foundation of a multilevel approach lies in the inexorable dependency of
30 individual's health on the surrounding social and physical environment. Social contexts and
31 their characteristics are integral determinants of variations in health outcomes both within
32 populations and between populations. Societal patterns of oral diseases represent the

33 biological consequences of living and working conditions differentially afforded to social
34 groups as a product of economic and political priorities of societies.^{3,9-11} These priorities vary
35 across different levels of administrative and political boundaries ranging from global and
36 national, to small areas like neighbourhoods, local areas, municipalities, performing a
37 fundamental role in the distribution and access to oral health promoting/risks factors.
38 Multilevel studies pay equal attention to both intimate and ultimate causes of health without
39 discounting that individual health and its individual risk factors do not function in isolation.⁹
40 Hence, the opportunities and challenges offered in multilevel approaches for a better
41 understanding of the role of societal determinants of oral health need to be highlighted.

42 Testing hypotheses at multiple levels of social organization using multilevel studies, although
43 very useful, are complex.¹² These complexities arise due to the mutual interactions and
44 interdependencies between individual-level and societal-level factors related to health.
45 Methodological experts in epidemiology, particularly social epidemiologists, have
46 highlighted some challenges (relevant groups/levels, lag times, fallacies, confounding among
47 others) in the light of general health outcomes.¹²⁻¹⁵ Oral health conditions provide unique
48 opportunities to study social inequalities in health.¹⁶ The aetiological factors for two
49 commonly prevalent oral diseases (dental caries and periodontal disease) include a complex
50 mix of biological, environmental, and social influences.¹⁶ Many high-income countries
51 continue to exclude dental health-care from their universal health coverage policies leaving
52 the responsibility on individuals to cover and manage dental expenses.¹⁷ Collectively, social
53 determinants across different levels of social organization may systematically determine the
54 magnitude of oral health inequalities both within and between societies. Consequently,
55 general health and oral health conditions often differ in strength and the type of their
56 relationship with social determinants when examined simultaneously within the same
57 context.^{18, 19} For example, social inequalities in oral health were found to be more
58 pronounced than those in general health in Canada.¹⁸ A stronger association between income
59 inequality and dentition status (presence of teeth) than between income inequality and self-
60 rated health was reported in Japan.¹⁹ Accordingly, multilevel studies on societal determinants
61 of oral conditions can benefit from attention to theoretical and methodological considerations,
62 some of which may uniquely apply to oral conditions.

63 In the light of the public health challenges presented in oral health at the population level, this
64 paper aims to:

- 65 i) present theoretical and pragmatic reasons to consider re-balancing the weight of
66 studies on individual-level variations in oral health towards studies on population-level
67 variations in oral health,
68 ii) discuss the contribution of multilevel studies to generate valuable evidence on societal
69 determinants of oral health and understanding population variations in oral health, and,
70 iii) review methodological aspects relevant to the application of multilevel studies in oral
71 health.

72 **The case for more studies on population variations in oral health and its determinants**

73 Geoffrey Rose's seminal work in 'The Strategies of Preventive Medicine' stressed the
74 distinction between the risks for individual variations in health, and, the risks for population
75 variations in health.²⁰ Rose's theorem' or the 'prevention paradox' states that "a large number
76 of people exposed to a small risk may generate more cases than a small number exposed to a
77 high risk". Therefore, when everyone is exposed to the risk within a population, it is not
78 possible to measure the effect of the exposure without reference to a population that has a
79 different level of exposure. Building on this, Rose established that the determinants of
80 variations in health between populations differ from the determinants of variations in health
81 within the population.²⁰ This is a key argument for studying population variations in health
82 and its determinants. Differences in the magnitude of the influence of determinants of caries
83 rates among children within and between Australia and Vietnam is one example of
84 operationalization of Rose's theorem in oral epidemiology.² The study reported that while
85 individual social position was relevant for individual risk, country-level economic
86 development and social inequality were more relevant for population risk.²

87 Another epidemiologist, McMichael²¹ raised a relevant question that further underscores a
88 key argument for conceptualizing and studying health outcomes at a population level that is
89 relevant to oral health:

90 "Are we, merely distinguishing between upstream social contexts and their downstream
91 proximate manifestations? Or is there a category of risk factor that, in some collective way,
92 influences the health of the population at large via processes that have no direct downstream
93 manifestation?"

94 Evidence summarised below from the oral health literature reinforces the need to pursue this
95 line of enquiry.

96 *Growing evidence on the independent contribution of contexts in shaping oral health: A*
97 *systematic review,*²² and a scoping review,²³ have separately confirmed poor oral health
98 outcomes to be associated with less favourable contextual socioeconomic variables (high
99 area-level social inequality, high area-level deprivation, low social capital, and, low access to
100 dental healthcare). Socioeconomic inequalities in oral health outcomes according to contexts
101 are expressed spatially based on the variations in area-level social, political, and economic
102 characteristics. Therefore, population-level variations in oral health outcomes can finely
103 capture the population-level impact of contextual societal disadvantage.

104 *Population variations in oral health reveal underlying societal determinants:* Investigating
105 patterns of variation in oral health between populations is important in its own right. The
106 observed population-level variations are important to understand the significance of specific
107 contexts for oral health outcomes.²⁴ Otherwise, similar individuals may have differences in
108 their health dependent on where they live because of differing cultural, economic, political,
109 geographic, climatic and historical contexts.²⁵ The more homogenous the oral health of
110 individuals within a population, the higher the probability that determinants of oral health are
111 directly related to the contextual environment or the population characteristics. Interventions
112 in this case needs to be directed to contexts rather than individuals.²⁶ From an equity
113 perspective, population-variations in health outcomes, that are systematic, socially produced
114 and unfair, are highly relevant.²⁷ These inequities result from systematic differences in
115 exposure to health risks and protective factors as well as to treatment services, based on
116 social position.²⁸ Therefore, the observation of population variations in oral health between
117 societies reflects the need to understand the differences in characteristics of these societies.

118 *Socio-political and multilevel nature of oral health determinants:* Individual-level risk factors
119 for most prevalent oral conditions include high sugar consumption, tobacco use, lack of
120 access to fluoride, high levels of stress, lack of oral hygiene and favorable pattern of dental
121 visiting.²⁹⁻³¹ The distribution of these individual-level determinants both within and between
122 societies can be influenced by societal determinants and policy decisions at multiple
123 geographic and administrative levels. Variations in presence and intensity of policy
124 implementation can also exist between the geographic and administrative levels. Key policy

125 determinants that impact distribution of individual-level behavioural risk factors include
126 federal, state and local level decisions on community water fluoridation;³² health care
127 arrangements including provision and financing; tobacco control policies including
128 ratification of Framework Convention of Tobacco Control (FCTC) at the national level to its
129 compliance at different sub-national geographic and administrative levels;³³ trade
130 arrangements/agreements and marketing regulations impacting food demand and supply;³⁴
131 and availability of local physical and social environments that improve social cohesion and
132 physical activity.³⁵ Studying population variations in oral health can allow comparisons
133 between societies and provide key insights about existing policies and the impact on
134 population oral health of different political and administrative arrangements. Cross-national
135 studies comparing countries with different policies, for example regarding taxation of sugar
136 foods/beverages or their dental care systems, can contribute significantly in assessing the
137 potential impact of upstream interventions on oral health. Additionally, natural experiments
138 at the societal level that compare population oral health can serve as a useful tool. Natural
139 experiments applied in oral health context from Brazil and Japan have improved the current
140 understanding of the effectiveness of water fluoridation in reducing dental caries among
141 adults, and the impact of socioeconomic disadvantage on tooth loss.^{36,37}

142 *Explanatory potential of individual-level studies for population oral health:* Most
143 epidemiological studies report measures of individual relative risk (odds ratio (OR), risk ratio
144 (RR) rather than the population attributable risk (PAR).³⁸ PAR describes how much of the
145 condition within the population can be attributed to a particular risk factor, while the risk
146 ratio (RR) informs the change in risk of an outcome among exposed individuals compared to
147 unexposed individuals. Even with larger and statistically significant levels of RR, the PAR
148 can be smaller and insignificant from a public health perspective, if the exposure is not
149 widespread.³⁸ Alternatively, a low RR can accompany a high PAR when an exposure occurs
150 frequently in the population. The study by Do² on the differences between caries rates among
151 Vietnamese and Australian children found an RR of 1.24 for dental caries among Vietnamese
152 children who did not start brushing with toothpaste before three years of age. The RR for
153 Australian children was similar with a value of 1.27. However, the Population Attributable
154 Fraction (PAF - the proportion of the disease in the population attributable to a factor of
155 interest) for Vietnamese children by introducing brushing with toothpaste before the age of
156 three years was 18% compared to only 3% for Australian children for the prevention of
157 caries. Lack of reporting of PAR along with measures of association in studies of individual-

158 level outcomes limits the knowledge of the preventive capacity of interventions for
159 population oral health.

160 Individual-level risk factors for oral diseases often do not vary enough within populations to
161 permit quantification of their probability to increase risk at an individual level.³⁹⁻⁴¹ This issue
162 further limits the value of individual-level studies in generating evidence for population-level
163 prevention, even when PAR is reported. For instance, the WHO recommends that free sugars
164 intake should be restricted to less than 10% of total energy. A conditional recommendation
165 for further health benefits particularly with regard to dental caries includes restriction to less
166 than 5% of total energy.⁴² This recommendation is exceeded in most countries. A review of
167 data on sugar intake from national surveys from Australia, Denmark, Ireland, Norway and US
168 showed that at a population level none of these countries met the recommendation of 5%
169 limit.⁴³ Furthermore, evidence from Australia demonstrates the prevalence of exceeding the
170 recommendations is high (52% for the 10% recommendation and 89% for the 5%
171 recommendation).⁴² Therefore, there may not be enough variation in the exposure within a
172 population to the effects on dental caries to be determined. Ecological studies, which study
173 between-population, rather than within-population variation in caries according to sugar
174 availability, report larger variations in caries status according to sugar availability when
175 compared to individual-level studies.⁴⁴ In cases where the individual risk factors do not vary
176 within populations, evidence on population variations in oral health are likely to be more
177 informative in making public health decisions.

178 *Informing strategies for prevention for oral diseases:* During the second half of the 19th and
179 first half of 20th century, there was a shift in epidemiology away from studying societal
180 causes of diseases and a move towards the individual and microbial causes.^{45, 46} and is
181 identified as an epistemological revolution in understanding the causes of diseases.⁴⁶
182 Different approaches to disease causality have important political and medical implications as
183 they mean a different locus of responsibility for prevention of diseases. A causal focus on
184 microbial factors confers responsibility of prevention to health professionals, individual
185 behaviours or lifestyle factors implies a personal responsibility for disease control, while a
186 socio-environmental causal model places responsibility on authorities and general society for
187 the prevention of disease and reduction of exposure.⁴⁷ Prevention strategies for non-
188 communicable diseases including oral diseases often suffer from a similar individually-
189 focussed approach by promoting change in individual risk-factors. The population-based
190 strategy, the high-risk strategy, and the directed population strategy are the three types of

191 strategies applied towards prevention of oral diseases and promoting oral health. The
192 population-based strategy for prevention starts with the recognition that the occurrence of
193 common diseases and exposures reflects the behavior and circumstances of society as a
194 whole.⁴¹ Alternatively, the high-risk strategy targets individuals identified as having an
195 elevated risk of some adverse health outcome.⁴¹ The directed population strategy is a version
196 of the population strategy but it is directed more towards vulnerable groups based on their
197 social circumstances rather than elevated levels of risk.^{48, 49} The studies on the causes of
198 individual variations in oral diseases generate evidence that may provide limited support to
199 whole population approach for prevention. For instance, Holst and colleagues have reported
200 that the occurrence of a carious lesion in individuals and the occurrence of caries in
201 populations have different causal candidates and patterns. This exemplifies the distinction
202 between the causes of cases and the causes of incidence in a population.^{50, 51} Individual-level
203 approaches have remained as the dominant paradigm in understanding the production and
204 prevention of oral diseases.^{4, 52} This approach is consistent with the ‘high-risk strategy’²⁰ and
205 has evolved from both the biomedical nature of dentistry and an individual ‘risk factor’ focus
206 from clinical oral epidemiology.⁴ The limitation of a ‘high-risk strategy’ in reducing
207 variations in population levels of oral health is well established within the literature.^{4, 53} This
208 approach does not acknowledge the growing understanding of the multilevel nature of health
209 determinants⁹ and societal determinants in shaping the distribution of oral health.^{4, 54, 55}
210 Therefore, dominance of individual based approaches shifts attention from underlying
211 societal determinants of health and encourages individual responsibility to maintain oral
212 health rather improving environments to promote oral health.⁵⁶

213 **Advancing the multilevel study approach**

214 An ecological design within epidemiology seeks to understand how contexts affect the health
215 of groups through selection, distribution, interaction, adaption and other responses.⁵⁷
216 Multilevel studies investigate both groups and individuals as the unit of analysis. It allows the
217 simultaneous investigation of between-group and within-group variability in individual-level
218 outcomes.¹² Therefore, multilevel studies can be applied to examine the associations between
219 group level and individual level variables with individual-level outcomes. Additionally, it can
220 be applied to examine between-group and within-group variability and the contributions of
221 group-level and individual-level variables to variability at both levels -population variations
222 in health and its determinants.⁵⁸

223 A key advantage of multilevel study is its potential to address confounding generated from
224 variables at alternate levels of social organizations when simultaneously analyzing variables
225 at two or more levels of social organization, multilevel studies allow addressing. This
226 advantage of multilevel studies has been widely exploited in studies of area-level income
227 inequality and health outcomes.⁵⁹ Early ecological studies on area-level income inequality
228 and population health using single-level regression models have been criticized in the past. It
229 is debated that the observed associations between area-level income inequality and average
230 health status at the population level in ecological studies were due to the effect of individual
231 income on individual health (compositional effect) rather than a true effect of income
232 inequality.⁶⁰ Multilevel studies offer the opportunity to separate the contextual effect of
233 income inequality on individual health from the compositional effect of individual income by
234 allowing to adjust for individual income within the same regression.⁵⁹ However, ecological
235 studies analyzed population risk according to area-level income inequality, while the
236 multilevel studies assessed individual risk according to area-level income inequality.⁶¹ The
237 population-level aspect of health outcome in multilevel studies is studied through
238 investigating the share of individual-level variation in health outcomes that exist at the
239 population level through decomposition of variance.⁶¹

240 Methodological experts argue that multilevel modelling has not been used to its potential to
241 answer questions on population-level variations in health status and its determinants in the
242 field of social epidemiology.^{24, 61-65} Studies have mostly focussed on average associations
243 between individual and societal determinants, and health outcomes, ignoring a thorough
244 analysis of heterogeneity around average associations examined through the variance
245 estimates obtained from multilevel studies.⁶¹ The variance component informs to what extent
246 individuals within a group are correlated with one another in relation to health. The extent of
247 clustering has value in the context of ideas about considering interventions on places instead
248 of people.²⁶ One application of this logic is identified in a study where multilevel modelling
249 is utilized to identify appropriate geographic levels for policy intervention.⁶⁶ Geographic
250 levels at which the observed variations in outcomes are larger, there may be greater potential
251 for policy intervention to have an impact on the outcomes of interest, compared with
252 targeting policy at levels with relatively smaller variations.⁶⁶ Multilevel studies also provide a
253 suite of measures based on average association between societal exposures and individual
254 health outcomes (OR, RR), and measures of variation in individual health (variance) and its
255 decomposition at the population level (variance partition coefficients (VPCs), intra-class

256 correlation coefficient (ICC) for continuous outcomes and median odds ratio (MOR) for
257 binary outcomes), that can be applied to understand societal causes of population variations
258 in oral health. Two additional measures: 80% Interval Odds Ratio (IOR) and Proportion of
259 Odds Ratio in Opposite Direction (POOR), can be quantified by combining regression
260 coefficients obtained from averaged associations between societal determinants and
261 individual oral health and the variance attributed to the contextual level. The two measures
262 estimate the heterogeneity in the associations between societal exposures and individual
263 health outcomes among contexts/population groups.⁶¹ Measures of variation in individual
264 health and its decomposition are critical for inferences on population-level variations in
265 health. In addition to ICC, measures of discriminatory accuracy such as Area Under Curve
266 (AUC) can be applied to understand the independent contribution of societal context in
267 general, and of specific societal exposure, in determining oral health outcomes.⁶¹
268 Collectively, these measures can be exploited to provide a thorough and realistic assessment
269 of the relationship between societal determinants and oral health within the same dataset.

270 Predominantly, multilevel studies on societal determinants of oral health are of two kinds.
271 Some studies have simultaneously examined the role of multiple societal determinants
272 (Human Development, access to fluoridated tap water, oral health coverage, and income
273 inequality) and oral health outcome/s consistent with a more exploratory approach using the
274 social determinant framework.⁶⁷⁻⁶⁹ Others have tested specific associations between one
275 societal determinant (for example area-level income inequality, neighbourhood
276 deprivation)^{19, 70-73} and oral health outcome/s consistent with a causal approach. The
277 dominance of probabilistic risk factor epidemiology has limited the use of multi-level models
278 to examine between-group and within-group variability through quantification of variance
279 and its decomposition at different levels of social organizations.^{64, 65} The understanding of the
280 social determinants oral health can substantially benefit from the application of multilevel
281 models by examining between-group variability in individual-level oral health outcomes as a
282 method to study population-level variations in oral health.

283 **Methodological aspects relevant to application of multilevel approaches within oral** 284 **health**

285 Methodological considerations related to multilevel studies relevant to oral health are collated
286 from the general health literature and discussed below under logical headings. Wherever

287 possible, published or hypothetical examples from oral health are used to illustrate their
288 relevance.

289 *Types of cross-level associations and arising fallacies*

290 The simultaneous assessment of associations between societal and individual factors and
291 individual health outcomes in multilevel studies has led to investigations of three main
292 different types of associations.¹⁵ A societal exposure can potentially impact oral health at an
293 individual level through direct cross-level association, indirect cross-level association, and
294 cross-level effect modification. A direct cross-level association occurs when a societal factor
295 has a direct impact on the individual oral health outcome. For example, a person living in an
296 area with community water fluoridation (exposed to fluoride) has lower risk of dental caries,
297 than a person in non-fluoridated area.³² Indirect cross-level association occurs when a societal
298 factor results in a change in individual-level exposure, which consequently, increases or
299 decreases risk of disease at an individual level. For instance, the presence of school policies
300 on the availability of sugar-sweetened beverages (SSBs) can discourage individual
301 consumption of during the day, therefore, reduce the risk of dental caries.⁷⁴ Finally, cross-
302 level effect modification occurs when a societal factor modifies the association between an
303 individual level factor and individual health outcome. Some evidence exists to suggest that
304 the associations between individual social position and oral health vary according to the
305 welfare typology,^{75, 76} in line with the cross-level effect modification. Clarity on these
306 pathways when generating hypothesis is critical as the findings have consequences of the
307 choice of policy intervention points for improving oral health.

308 Several fallacies are produced in a situation when the hypothesis generated in both
309 conventional ecological studies and multilevel studies are not theoretically aligned with the
310 potential mechanisms of how societal factors can impact oral health. These fallacies are
311 called ecological, atomistic, sociologicistic, and psychologistic, and are widely discussed in
312 general health literature.^{15, 77} Each of these fallacies are described along with a suitable
313 published or hypothetical example in Table 1.

314 *Ecological variables: classification and constructs*

315 Ecological variables represent group-level properties, including societal factors, which are
316 relevant to oral health. Depending on their measurement or the construct they aim to

317 capture, ecological variables have been classified in several ways within the literature.
318 Classification of ecological variables reveals its degree and nature of dependency on
319 individual-level factors. For instance, ecological variables can be integral or derived.¹²
320 Integral ecological variables are only group characteristics, and cannot be measured at an
321 individual level, for example community water fluoridation and air pollution. Conversely,
322 derived ecological variables present as a mathematical summary of individual
323 characteristics within a group,¹² for example percentage of children with sugar
324 consumption above the World Health Organization recommendation, or area-level mean
325 income. However, derived ecological variables may or may not have their individual-level
326 analogue. While area-level mean income has an individual income as its individual
327 equivalent, area-level income inequality is solely a group property and does not have an
328 individual equivalent.

329 Based on the constructs they capture; ecological variables can be categorized as:

- 330 i. aggregate/ contextual/ analytical,
- 331 ii. contagion,
- 332 iii. environmental,
- 333 iv. structural, and
- 334 v. global/ integral.

335 The description of these categories of ecological variables along with suitable examples is
336 presented in Table 2. Clarifying the constructs that the ecological variable of interest aims to
337 capture has implications on measurement issues and analytical approaches. For instance,
338 global variables such as legislations and policies are likely to have a more diffused effect
339 among populations rather than leading to an instant biological or bio-behavioural impact on
340 ‘high-risk’ individuals. In such cases ecologic inferences about effects on group rates or
341 population-level variations may be more relevant than individual risks.⁴⁰

342 *Meaningful population groups, scale, and unique characteristics*

343 Specifying meaningful boundaries and identifying groups of interest for the ecological unit of
344 interest is core to any multilevel study.^{14, 78} Despite the use of ‘population’ across many
345 disciplines analysing population data—for example, epidemiology, demography, sociology,
346 ecology, population biology and population genetics, statistics, and biostatistics, it is rarely
347 defined, except in abstract statistical terms.⁷⁹ Various criteria can be applied to define
348 population groups of interest. For instance, the boundaries of a ‘neighbourhood’ can be

349 defined based on historical or geographic criteria, the perception of the residents or the
350 administrative boundaries used for policy delivery. Moreover, 'neighbourhood',
351 'community', and 'area' are often used loosely within the health literature to identify an
352 individual's immediate residential environment, and the three terms are not explicitly defined
353 or distinguished.⁸⁰ The population-level effectiveness of public policies such as community
354 water fluoridation in reducing dental caries are more consistent with administratively defined
355 boundaries, compared to interventions to improve opportunities for social interactions.
356 Creating opportunities for social interaction in a community is likely dependent on what an
357 individual perceives as the boundary for a community rather than the administratively defined
358 limits. Recently, it was highlighted that a "residential" effect fallacy bias exists in most
359 studies of neighbourhood and health studies that ignorantly capture non-residential
360 environment effects, leading to overestimation of residential intervention effects.⁸¹ These
361 non-residential environment effects may be due to schools or workplaces depending on the
362 health outcome, population density, and individual mobility.^{8, 81}

363 The selection of spatial scale for testing associations between ecological factors and health
364 outcomes is both an important theoretical and methodological aspect. First, the societal
365 processes that produce health may vary by geographic scale.⁸² Second, group-level
366 characteristics do not occur randomly and are based on the social and political context that
367 influence these characteristics. The spatial scale of assessment has been used consistently as
368 one of the most important explanations for the lack of association income inequality and
369 general health outcomes at a sub-national and/or small area level.⁸³ Studies have examined
370 associations between income inequality and health outcome at different levels of aggregation
371 within the same country and found significant variations.⁸⁴ The lack of association at a
372 smaller level of geographic aggregation and the presence at the larger is attributed to the
373 inability of income inequality as an exposure to reflect the social stratification within a
374 society at a small area level.⁸³ Medical geographers have also recognized the 'modifiable
375 areal unit problem (MAUP)' and 'uncertain geographic context problem (UGCoP)' that need
376 to be considered when selecting the relevant spatial scale.⁸⁵⁻⁸⁸ MAUP relates to the fact that
377 societal exposures vary based on the definition of the geographic scale selected as well as
378 zonation areas even when one scale is selected.⁸⁵ Consequently, there is a possibility of
379 spatial misclassification of exposure, and the likelihood of a spurious association between
380 area-level factors and oral health outcome.^{85, 86} Consistent with MAUP, exposure
381 misclassification based on the selection of neighbourhood definition has been empirically

382 shown for the exposure of youths' access to tobacco retailers in a study.⁸⁵ UGCoP identifies
383 two sources of contextual uncertainty. These sources include spatial configuration of
384 geographically defined contexts and the timing and duration of exposure to those contexts.⁸⁷
385 ⁸⁸ However, the role of spatial aggregation and individual mobility has not been dealt in
386 multilevel studies of oral health.

387 Explicit definitions of ecological factors are crucial when generating hypothesis on societal
388 determinants of health. This applies also to the clarity on levels (societal or individual) at
389 which ecological factors are measured. The level of measurement has consequences on
390 theoretical pathways through which they impact oral health outcomes. Differences in
391 definitions of concepts might exist according to levels. For instance, there is a lack of
392 consensus on the meaning and definition of social capital.⁸⁹ Lack of clarity on the definition
393 makes the operationalization of social capital in epidemiological investigations challenging.
394 Social capital is a contextual construct– a societal property. However, social capital is often
395 measured at a societal level through deriving aggregates of individuals' perceptions of
396 reciprocity, trust, and, engagements in civic activities. Social interactions among residents are
397 rarely captured at the contextual level.⁹⁰ Individual perceptions of contextual social capital
398 may potentially vary within the same context. Therefore, relying on aggregated measures of
399 social capital that are unadjusted for individual-level variations in perceptions can lead to
400 potential misclassification. This complexity in the measurement of social capital reflects the
401 need for the explicit meaning of ecological measures at the contextual level. Additionally,
402 recognizing the diversity of multiple mediating pathways (social capital or neo-material
403 factors) for each and every oral health outcome and, at both individual and population levels
404 can be helpful in a better understanding of causal relations and potential interventions.²³
405 While social capital explained the association between income inequality and self-rated
406 health in Japanese adults, it did not explain their dentition status.¹⁹

407 Most multilevel studies are a secondary analysis of already collected data. Consequently,
408 researchers may be forced to use imperfect proxies for measuring group level constructs. This
409 provides limited information and can further make inferences drawn from such analysis
410 inaccurate.⁸ Caution is required particularly in identifying appropriate population groups,
411 spatial scales and differentiating between the unique properties of ecological factors in
412 interest.

413 *Role of lag times*

414 Failure to recognize and account for lag time between an ecological exposure and individual
415 health outcome is a form of misclassification bias. The role of lag times between exposure
416 and outcome has been paid less attention than other challenges in multilevel studies.¹⁵
417 Usually, multilevel studies are conducted using cross-sectional data where the distinction
418 between current and past exposures cannot be made. Societal factors are not likely to have an
419 instantaneous effect on individual health, and therefore establishing appropriate lag period
420 between the exposure and specific oral health outcome is necessary particularly when the
421 exposure is not stable over time.¹⁵ The lack of association between a societal exposure and an
422 oral health outcome due to the inappropriate definition of lag times can be misleading as
423 associations may be present when appropriate lag times are considered. Therefore,
424 assessment of lag time is critical before dismissing the evidence on the impact of societal
425 determinants on oral health based on findings where the exposure is non-stable and exposure
426 and outcome are measured simultaneously.

427 *Current challenges and limitations with multilevel statistical modelling*

428 Multilevel models are still evolving. Issues such as appropriate sample size, methods for
429 selecting and reporting appropriate measures of interest, and the reporting of diagnostic tests
430 within multilevel studies are yet to be resolved.¹⁴ Model diagnostics are also seldom reported
431 within studies.¹⁴ Assumptions of multilevel modelling regarding the hierarchical units being
432 independent of each other are also rarely met. A lack of reporting of measures of variation in
433 individual health and its decomposition is also identified within the literature.⁶⁵

434 A more conceptual issue relevant to oral health needs further examination in the application
435 of multilevel models in studying population variations in oral health. Compared to general
436 health outcomes like mortality and health that are captured widely in census data and
437 registration data, for oral health information data is obtained from oral health surveys that are
438 not designed with a primary purpose to make inferences at smaller geographies, and are
439 underpowered for this purpose. This limits the examination of average associations between
440 an area-level societal determinant (area-level income inequality, area-level deprivation) and
441 population oral health (rates of dental caries, rates of oral cancer) at small area level in
442 multilevel studies. This restricts the assessment of theoretical pathways proposed to explain
443 population-level variations in health/disease rates according to societal determinant when

444 applied to explaining individual-level variations in oral health/disease. Some of the mediating
445 pathways operate more strongly at an environmental level (legislation, policies, social capital,
446 access to health services) while others at the individual level (stress, health behaviour,
447 utilization of health services). Therefore, theoretical pathways need to be proposed and
448 defended based on the level at which each oral health outcome is analysed. Potential
449 differences in strengths of associations at the population level (population risk), and at the
450 individual level (individual risk), may also demand separate prevention strategies and policy
451 responses.⁹¹ The extent to which studying population variations in oral health in multilevel
452 studies through analysing variance share at population level informs these two policy-
453 relevant issues needs further assessment.

454 Power and sample size calculations for multilevel hypotheses are complex as power depends
455 both on a number of groups as well as the number of individuals per group.¹³ Calculation of
456 sample size in multilevel studies is dependent on the level at which inferences are to be made.
457 When these are at the group level, there should be a sufficient number of groups rather than
458 individuals. But, when the inference is to be made at the individual level, then both sufficient
459 number of groups and individuals are required. Often multilevel studies are challenged due to
460 the small size of groups. Simulation studies have shown that multilevel models with large
461 numbers of groups (more than 459 groups) even with smaller group sizes remain stable, and
462 neither fixed or random components are affected due to group sizes.⁹² Since most multilevel
463 studies on societal determinants of oral health use secondary data, Monte Carlo simulation of
464 the model should be applied to estimation post-hoc power and for sample size calculation.⁹³
465 Most multilevel studies analyse cross-sectional data where the temporal sequence between
466 exposure and outcomes cannot be established. Multilevel studies on longitudinal datasets can
467 help resolve this issue as temporal sequence between the societal exposure and oral health
468 outcomes can be established. However, multilevel statistical modelling is mainly applied in
469 longitudinal data to manage data imbalances due to loss to follow up, rather than to examine
470 associations between societal determinants and oral health outcomes.

471 **Final remarks**

472 The challenges currently posed in population oral health highlight the need for more
473 population focussed research and the use of ecological studies in the field or dental public
474 health. The value in studying population-variations in oral health and its determinants has a
475 rationale embedded in theory and is fundamental for policy assistance. This will likely

476 contribute towards a better understanding of how exposures that affect all individuals in a
477 population contribute to their oral health. There is a need for balancing the weight of
478 individual-level studies with studies of population variations and societal determinants, not to
479 replace the individual-level studies, but to complement them.

480 Ecological studies offer an opportunity to study average associations between societal
481 determinants and population-level variations in oral health, but cannot account for potential
482 confounding introduced by factors from alternate levels of social organization.⁶ Multilevel
483 studies using individual and societal data collectively, overcome this limitation by
484 simultaneously examining multiple hypotheses generated at different levels of social
485 organization. Using multilevel models to quantify the share of individual-level variation in
486 oral health outcomes that exist at a societal level, the contribution of societal and individual
487 determinants on this share of variance, allows the investigation of population-level variations
488 in oral health and its determinants.⁶¹

489 Multilevel studies of societal determinants of oral health require careful attention from the
490 stage of conceptualization to design, analysis and reporting, as highlighted in this paper.
491 These features are not unique to such studies and form the basis of any scientific enquiry. In
492 addition to multilevel methods, studies on societal determinants of oral health can deal with
493 inherent complexity by exploring methodological approaches from other disciplines such as
494 social and political sciences including qualitative methods. Finally, studies with explicit
495 theoretical bases²³ that draw on the strengths of multilevel modelling can provide a more
496 enhanced understanding of societal determinants of oral health, and consequently lead to
497 robust evidence for relevant policy solutions.

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Table 1. Description of fallacies along with suitable examples from oral health

Fallacy	Reason	Description	Example
Ecological	Construct and measurement issues	Associations at ecological level are used to make inferences on the association at an individual level due to absence of data at an individual level. The more heterogeneous the population, the higher is the fallacy	Association between water fluoridation and skeletal fractures ⁹⁴ : Supportive evidence for the association came largely from ecological studies comparing rates of fracture between fluoridated and non-fluoridated communities. However, well designed studies that measured individual exposure to water fluoridation/fluoride intake and controlled for different confounders could not find an association between dentally optimal doses of fluoride and fracture. This indicates a case when ecological level associations were not held true at the individual level.
Atomistic	Construct and measurement issues	Associations at individual level are used to make inferences on the association at an ecological level due to absence of data at a population level. This fallacy ignores the fact that societal factors and population has independent characteristics	Individual income may be negatively associated with tooth loss and it is inferred that mean income of an area is associated positively with population rate of tooth loss. However, the mean income may not be associated or positively associated with population rate of tooth loss.
Sociologicistic	Ignorance of variables from	This fallacy is a consequence of ignoring the role of individual level factors in group level	Effects of fluoride intake on population-level differences in dental caries is determined by testing correlations between

	individual level	associations	community-level water fluoridation and community levels of dental caries. Interpreting that community water fluoridation reduces every residents' risk of dental caries within such studies can be prone to sociologic fallacy as certain sub-groups may have preferences of bottled water over tap water.
Psychologicistic	Ignorance of variables from population level	This fallacy is a result of ignoring the role of ecological level factors in individual level associations	Ignoring the fact that water fluoridation is an environmental factor, and its presence may modify the association between fluoride intake and dental caries at the individual level.

Table 2. Description of different categories of ecological variables according to their classification and examples

Category	Description	Example
Aggregate/contextual/analytical	Aggregate summary measure of individual characteristics in a group (similar to derived variables)	Area-level mean income
Contagion	Aggregates of individual outcomes	Prevalence of dental caries and tooth loss rates of a group

Environmental	Physical characteristics with individual analogue	Environmental measure: Residential access to water fluoridation Individual analogue: Consuming fluoridated tap water
Structural	Patterns of relationship between individuals of a group	Social capital, social cohesion, social inequality as a product of power relations
Global	Attributes belonging to groups and not reduced to individuals	Legislations and policies



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