

Distinguishing between enamel fluorosis and other enamel defects in permanent teeth of children

Aira Sabokseir, Ali Golkari, Aubrey Sheiham

Background: The inconsistent prevalence of fluorosis for a given level of fluoride in drinking water suggests developmental defects of enamel (DDEs) other than fluorosis were being misdiagnosed as fluorosis. The imprecise definition and subjective perception of fluorosis indices could result in misdiagnosis of dental fluorosis. This study was conducted to distinguish genuine fluorosis from fluorosis-resembling defects that could have adverse health-related events as a cause using Early Childhood Events Life-grid method (ECEL).

Methods: A study was conducted on 400 9-year-old children from areas with high, optimal and low levels of fluoride in the drinking water of Fars province, Iran. Fluorosis cases were diagnosed on the standardized one view photographs of the anterior teeth using Dean's and TF (Thylstrup and Fejerskov) Indices by calibrated dentists. Agreements between examiners were tested. Early childhood health-related data collected retrospectively by ECEL method were matched with the position of enamel defects. **Results:** Using both Dean and TF indices three out of four dentists diagnosed that 31.3% (115) children had fluorosis, 58.0%, 29.1%, and 10.0% in high (2.12-2.85ppm), optimal (0.62-1.22ppm), and low (0.24-0.29ppm) fluoride areas respectively ($p < 0.001$). After matching health-related events in the 115 (31.3%) of children diagnosed with fluorosis, 31 (8.4%) of children had fluorosis which could be matched with their adverse health-related events. That suggests that what was diagnosed as fluorosis were non-fluoride related DDEs that resemble fluorosis. **Discussion:** The frequently used measures of fluorosis appear to overscore fluorosis. Use of ECEL method to consider health related events relevant to DDEs could help to differentiate between genuine fluorosis and fluorosis-resembling defects.

1 **Distinguishing between enamel fluorosis and other enamel defects in permanent**
2 **teeth of children**

3 Aira Sabokseir¹, Ali Golkari^{1*}, Aubrey Sheiham²

4 ¹ Department of Dental Public Health, School of Dentistry, Shiraz University of
5 Medical Sciences, Shiraz, Iran,

6 ² Department of Epidemiology and Public Health, University College London,
7 London, UK.

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9 ***Corresponding author:** Ali Golkari, Shiraz School of Dentistry, Ghomabad,
10 Ghasrodasht St. Shiraz, Iran. Phone: +98 9175607254 Fax: +98 7136270325 e-mail:
11 golkaria@sums.ac.ir

13 **Abstract**

14 **Background:** The inconsistent prevalence of fluorosis for a given level of fluoride in
15 drinking water suggests developmental defects of enamel (DDEs) other than fluorosis
16 were being misdiagnosed as fluorosis. The imprecise definition and subjective
17 perception of fluorosis indices could result in misdiagnosis of dental fluorosis. This study
18 was conducted to distinguish genuine fluorosis from fluorosis-resembling defects that
19 could have adverse health-related events as a cause using Early Childhood Events Life-
20 grid method (ECEL).

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22 optimal and low levels of fluoride in the drinking water of Fars province, Iran. Fluorosis
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24 using Dean's and TF (Thylstrup and Fejerskov) Indices by calibrated dentists.
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26 collected retrospectively by ECEL method were matched with the position of enamel
27 defects.

28 **Results:** Using both Dean and TF indices three out of four dentists diagnosed that
29 31.3% (115) children had fluorosis, 58.0%, 29.1%, and 10.0% in high (2.12-2.85ppm),
30 optimal (0.62-1.22ppm), and low (0.24-0.29ppm) fluoride areas respectively ($p < 0.001$).
31 After matching health-related events in the 115 (31.3%) of children diagnosed with
32 fluorosis, 31 (8.4%) of children had fluorosis which could be matched with their adverse
33 health-related events. That suggests that what was diagnosed as fluorosis were non-
34 fluoride related DDEs that resemble fluorosis.

35 **Discussion:** The frequently used measures of fluorosis appear to overscore fluorosis.
36 Use of ECEL method to consider health related events relevant to DDEs could help to
37 differentiate between genuine fluorosis and fluorosis-resembling defects.

39 Introduction:

40 Despite the extensive use of well documented indices of dental fluorosis (Dean 1942;
41 Fejerskov 1988), there is inconsistency in the reports on the prevalence rates of
42 fluorosis for a given level of fluoride in drinking water. Most probably, this inconsistency
43 in the prevalence of fluorosis occurs due to subjective perception of fluorosis by
44 examiners. Therefore, there is a strong possibility that other Developmental Defects of
45 Enamel (DDE) rather than excess intake of fluoride are being misdiagnosed as fluorosis
46 (Atar & Körperich 2010).

47 The two widely used indices of dental fluorosis are Dean's Index (Dean 1934; Dean
48 1942) and the Thylstrup and Fejerskov Index (TF Index or TFI) (Thylstrup & Fejerskov
49 1978; Fejerskov 1988). None of them clearly distinguish between defects caused by
50 fluorosis and caused by other factors. The differences between some of the diagnostic
51 categories are uncertain, vague, or insensitive (a clear example is the "questionable"
52 category in Dean's Index) . In Dean's Index, each individual is given one score, as a
53 score for the whole mouth, according to the two teeth most affected by fluorosis. This
54 index categorizes each tooth as normal, questionable, very mild, mild, moderate,
55 moderately severe, and severe. The classification is based on color and extent of
56 discolored enamel together with added hypoplasia in case tooth belongs to the last two
57 categories.

58

59 Thylstrup and Fejerskov reformulated Dean' Index as the TF Index (TFI). They used
60 both clinical and histological appearance of fluorosis and created a single coded index
61 from 0 (normal) to 9 (Thylstrup & Fejerskov 1978). The TFI was modified and finalized
62 in 1988 (Fejerskov 1988). From 1998, the scoring of facial surface was recommended
63 for TFI. Teeth should be cleaned and dried before examination. Cleaning and drying of
64 teeth make the appearance of fluorotic change more prominent and also the diagnosis
65 of questionable cases easier. Still the difference between some categories of TFI is not
66 clear.

67 As the main indices for diagnosing dental fluorosis include definitions that are imprecise,
68 it is not surprising that in some studies the reported prevalence of fluorosis was similar
69 despite the levels of fluoride in the drinking water being similar or different. For example,
70 a study in KwaNdebele (Africa) revealed that the prevalence of fluorosis was similar in
71 residents of areas with considerably different levels of fluoride in the water supply
72 (Lewis & Chikte 1995). A study in Andhra Pradesh (India) of four different areas with
73 different levels of fluoride in drinking water (< 0.7, 0.7-1.2, 1.3-4.0, >4 ppm) reported
74 100% dental fluorosis even in areas with optimum level of fluoride (Sudhir et al. 2009).
75 In Hong Kong, where the fluoride concentration of public water supplies was increased
76 from 0.5 ppm to 0.7 ppm, and then 1 ppm, the prevalence of DDE decreased
77 significantly from 92%, to 55%, and then 35% (Wong et al. 2006).

78 DDEs can be localized or generalized. Numerous systemic risk factors cause
79 generalized or "diffuse" DDEs (Small & Murray 1978; Pindborg 1982; Atar & Körperich
80 2010). The generalized DDEs may be of genetic origin (Thesleff 2000; Atar & Körperich
81 2010) or caused by malnutrition or diseases that occurred during early childhood
82 (Pindborg 1982; Atar & Körperich 2010). Fluorosis and some other generalized DDEs
83 are similar in appearance (Small & Murray 1978; Pindborg 1982). The inconsistent
84 reports on the prevalence of fluorosis suggest that non-fluoride related DDEs were
85 being misdiagnosed as fluorosis. Using teeth as markers of significant health related
86 events and considering these risk factors could help to distinguish between genuine
87 fluorosis and non-fluoride related DDEs that resemble fluorosis.

88 Genuine fluorosis can best be distinguished from other DDEs by relating a DDE to
89 particular life events, such as a significant health related event. Such events can be
90 reasonably recorded using the life-grid method that helps people remember past events
91 more accurately (Blane 1996). The life-grid method has been widely used and been
92 very successful in obtaining the precise timing of past events in both qualitative and
93 quantitative studies (Berney & Blane 1997; Holland et al. 2000; Bell 2005). A specially
94 designed life-grid for early childhood containing developmental milestones and shorter

95 periods of time for the earlier years, is the Early Childhood Events Life-grid method
96 (ECEL) (Golkari 2009).

97 As there is a need for accurate data of dental fluorosis, the inconsistency of reports on
98 the prevalence of fluorosis suggests that more precise definitions and diagnostic
99 methods are needed for diagnosing dental fluorosis and distinguishing enamel fluorosis
100 from other DDEs. Therefore a study was planned with the objective of distinguishing
101 genuine fluorosis from fluorosis-resembling defects that could have adverse health-
102 related events as a cause, instead of just excess fluoride intake.

103

104 **Materials and methods:**

105 This cross-sectional study was performed on 400 9-year-old children of Fars province,
106 Iran. The children were randomly selected from areas of Iran with high, optimal and low
107 levels of fluoride in the drinking water. Fluorosis cases were diagnosed using Dean's
108 Index and TF Index (Dean 1942; Fejerskov 1988) by calibrated dentists. To differentiate
109 between genuine fluorosis and fluorosis resembling defects, early childhood data were
110 collected retrospectively by the ECEL method (Golkari 2009).

111 Ethical permission was obtained from the Ethical Committee of Shiraz University of
112 Medical Sciences (SUMS) and the Educational Head Office of the Fars Province. A
113 written consent form explaining the objectives and the stages of the study were sent to
114 the parents of the selected children. They were included only if the parents agreed to
115 take part.

116 The level of fluoride in water of each town of the province was obtained from their
117 primary health care trust. Towns were categorized into one of three fluoride categories;
118 high, optimal, and low fluoride. One town was selected randomly from each category.
119 The selected towns were Gerash with high fluoride (2.12- 2.85 ppm) in water, Sepidan

120 with low fluoride (0.24- 0.29 ppm) in water, and Shiraz was the city with an optimal
121 range of fluoride (0.62- 1.22 ppm) in water .

122 Sample size was calculated based on the estimated prevalence of fluorosis in Iran
123 (61%), $d=6.1\%$ (10% expected prevalence), $\alpha=0.05$. As a result, a sample size of 246
124 was needed if a simple randomized sampling was used in all areas. The selection of
125 children in Gerash and Sepidan (with high and low fluoride levels) was done by simple
126 randomized sampling. However, in Shiraz (chosen as area with optimum fluoride level),
127 the simple randomization was not possible as it was a big city. Therefore, a stratified
128 randomization method had to be adopted. It was done using the four different
129 educational zones of the city. The number of required sample in this area was multiplied
130 by 2 ($k=2$) to increase the accuracy in the stratified sampling method that was used. The
131 response rate was suggested to be around 80%. Therefore, at the end, 100 9-year-old
132 schoolchildren of Gerash, 100 from Sepidan and 200 from Shiraz were selected from
133 lists of students obtained from Educational Head Office of each town.

134 Children aged from 9 years, 0 month to 9 years and 11 months who returned the signed
135 consent form were included in the study. Children were excluded if they had lived for
136 more than six months from birth to 5 years of age in other towns (determined during
137 interview with parents). Those who had less than 7 permanent incisor teeth, those with
138 orthodontic brackets, overlapping teeth, and large restorations or severe extrinsic stains
139 on their incisors were also excluded (determined during intra-oral examination).

140 Intra-oral examinations were carried out to select children conforming to the study
141 criteria. The examinations were performed using a headlight, disposable mirrors and
142 tongue blades with children seated on a chair. Photographs of the dentition of selected
143 children were taken for the diagnosis of dental fluorosis. A one-view photograph was
144 taken of the anterior part of dentition using a digital camera (Nikon D7100, AF-S VR
145 Micro-Nikkor 105mm f/ 2.8 G IF-ED) based on methods described by Wong et al.
146 (2005). Children were asked to close their incisor teeth edge to edge. Cheeks and lips
147 were retracted so that all anterior teeth and some parts of upper and lower gums were
148 visible. The camera was adjusted to 15 degrees above the perpendicular to the central

149 incisors' plane to minimize specula reflection and burn outs (Ellwood et al. 1996).
150 Immediately after taking each photograph, it was assessed to confirm its quality, and
151 was repeated if necessary.

152 During the fluorosis assessment phase, first, the photographs were randomly ordered to
153 prevent bias induced by the assessors' foreknowledge of the fluoride in the area.
154 Photographs were then assessed by eight calibrated dentists who were blind to the
155 clinical condition and town of residence of the subjects. Four calibrated dentists
156 assessed the photographs based on Dean's Index, and another four used the TF Index
157 (Dean 1942; Fejerskov 1988). All dentists observed the photographs on one computer
158 with identical settings. The diagnosis of fluorosis was confirmed only if three out of four
159 dentists of each group agreed.

160 The objective of the next stage of study was to obtain data on early childhood adverse
161 events. The parents of all children participating in the study were invited to school for
162 interview. Name and date of birth of each child were double checked with their parents.
163 Parents were interviewed using the ECEL method (Golkari 2009). Information regarding
164 gestational age (preterm, term, delayed), birth weight, number of births, type of delivery
165 (natural, caesarean or facilitated delivery), trauma to baby during birth, and newborn
166 vitality score was obtained. Questions were asked relating to personal life line,
167 residential status, occupation of parents/guardians, and child activity line which could
168 help parents to remember adverse health-related events. Afterwards, any illnesses the
169 child had suffered were recorded. For each illness the parent was asked about the
170 name or description of illness, age at which the illness started, duration, perception of
171 severity (mild, moderate, or severe), if went to doctor, medication if used, and
172 hospitalization. Information regarding hospitalizations (age, duration, reason, type of
173 anesthesia if used, and name of hospital), and falls and accidents (age, cause, trauma
174 to face or teeth, hospitalization, and breathing status right after accident as an indicator
175 of severity well known by parents) were also obtained.

176 The timing of childhood adverse health-related conditions was matched with the timing
177 of formation and calcification of each part of permanent incisors (Golkari 2009). If no

178 adverse life condition could be matched to the position of a defect diagnosed as
179 fluorosis, the case was considered as genuine fluorosis. However, if a health-related
180 adverse condition could be matched to an enamel defect diagnosed as fluorosis, the
181 case was considered as fluorosis-resembling defect. Using this method, diagnosed
182 fluorosis defects were divided into genuine fluorosis and fluorosis resembling defects.

183 SPSS software (version 22) was used for data analysis. Agreements between
184 examiners who assessed fluorosis using Dean's Index and TF Index were tested using
185 Kappa-coefficient. McNemar, and Pearson correlation tests were also used to compare
186 the results reported by each pair of dentists. The prevalence of fluorosis in the three
187 selected areas was compared, before and after adjustments for sex, by chi-square test
188 and logistic regression. Childhood adverse health-related conditions were matched to
189 enamel defects diagnosed as fluorosis one by one.

190

191 **Results:**

192 The number of 9-year-old children included in the study was 376; 171 (46%) girls and
193 196 (53 %) boys. The number of included children from Gerash (high F), Shiraz (optimal
194 F) and Sepidan (low F) were 88, 189, and 90 respectively .

195 Using both Dean and TF indices, three out of four dentists diagnosed that 31.3% (115)
196 of children had fluorosis. The percentage of fluorosis cases in areas with high, optimal,
197 and low range of fluoride in water was 58.0%, 29.1%, and 10.0% respectively
198 ($p < 0.001$). Logistic regression showed that there was a positive relationship between
199 fluorosis and fluoride in the drinking water ($p < 0.001$). There was no relationship
200 between fluorosis and children's sex ($p = 0.228$).

201 There were significant differences among dentists who scored photographs using
202 Dean's and TF indices. Among the four dentists who assessed photographs according
203 to Dean's Index, the difference in the number of cases diagnosed as fluorosis was
204 statistically different between each two dentists ($p < 0.001$). There was only a slight

205 (kappa was between 0 and 0.2) or fair (kappa was between 0.2 and 0.4) agreement
206 between them (Table 1). Similar results of agreement were observed among the four
207 dentists who scored children using the TF Index. Although there was not a high
208 agreement among dentists, a positive correlation was observed ($p < 0.001$) (Table 2).

209 Adverse past childhood health-related events were found in 311 (84.7%) of children
210 using the ECEL method. Fluorosis defects were found in 115 (31.3%) of children. The
211 timing of adverse health-related events could be matched with the position of what was
212 diagnosed as fluorosis in 31 (8.4%) of children. These were regarded as fluorosis
213 resembling defects. Therefore, it was concluded that 26.9% of what was first diagnosed
214 as fluorosis was in fact fluorosis resembling defects.

215

216 The overall percentage of genuine fluorosis was 22.9%: 47.7%, 20.6%, and 3.3% in
217 areas with high, optimal, and low fluoride areas respectively ($p < 0.001$). The percentage
218 of fluorosis resembling defects in areas with high, optimal, and low range of fluoride in
219 water was 10.2%, 8.5%, 6.7% respectively. The difference in percentage of fluorosis
220 resembling defects among the three areas was not statistically significant (Table 3).

221

222 **Discussion:**

223 Fluoride is one of the most successful measures for prevention of dental caries in public
224 health (Petersen & Lennon 2004). However, there has always been controversy about
225 using fluoride because of fluorosis (Sapolsky 1968; Null & Feldman 2003; Ananian et al.
226 2005). Reports of a high prevalence of fluorosis in communities have led to objections
227 to fluoride. Therefore, there is a need for a precise way to diagnose dental fluorosis.
228 Many local and systemic risk factors cause DDEs. Some non-fluoride related DDEs are
229 similar to enamel fluorosis and should be differentiated from genuine fluorosis. No
230 standard method has been established to differentiate them from one another. The main

231 objective of this study was therefore, to try a method to distinguish fluorosis from other
232 kinds of DDEs that look like fluorosis but are caused by adverse health-related events,
233 not by excess fluoride intake.

234 A systematic review reported that in Iran with an average concentration of fluoride in
235 water of 0.43 ± 0.17 ppm, the prevalence of fluorosis was 61% (Azami-Aghdash et al.
236 2013). Based on the level of fluoride, the reported prevalence of fluorosis was high and
237 questionable. The inconsistency of the prevalence of fluorosis with the level of fluoride
238 in water reported in that study and in many other different parts of the world (Lo &
239 Bagramian 1996; Sudhir et al. 2009; Arif et al. 2013) suggests that dental fluorosis was
240 misdiagnosed.

241 The overall prevalence of diagnosed dental fluorosis in the current study was 31.3%.
242 However, by using the ECEL method and considering health-related events, the
243 prevalence of genuine dental fluorosis was 22.6% as there were 8.4% with fluorosis
244 resembling defects. That illustrates that fluorosis could be ruled out as the main cause
245 of about 27% of what was incorrectly diagnosed as fluorosis.

246 The current study also showed significant differences among the dentists who scored
247 photographs to diagnose fluorosis according to Dean's and TF Indices ($p < 0.001$). This
248 finding indicates that both Dean and TF Indices are too subjective and therefore not
249 precise ways to diagnose dental fluorosis.

250 These two Indices could lead to misdiagnosis of fluorosis resembling defects as
251 genuine fluorosis. Tavener et al also concluded that interpretation of criteria could be
252 different among examiners and stressed the necessity for standard methods to measure
253 dental fluorosis (Tavener et al. 2007). Some studies have shown good to excellent
254 agreement among examiners (Kumar et al. 2000). However, comparing the methods,
255 the lack of bias was an advantage of the current study, as photographs were assessed
256 instead of clinical examinations.

257 This study indicates that by considering adverse health-related events, it is possible to
258 distinguish genuine fluorosis from fluorosis-resembling defects. When the timing of an
259 adverse condition matches the timing of development of the part of enamel defect, the
260 adverse event could be the cause of the defect, not fluoride. If no adverse condition
261 could be matched to a defect, excess fluoride could, with caution, be considered as the
262 cause. In case of generalized defects, fluoride, genetic, or severe underlying systematic
263 disease, either individually or as a combination could be the cause.

264 One limitation of this study is that even if an adverse health-related event could be
265 exactly matched to a fluoride-resembling defect in terms of time and place, it could not
266 be definitely considered that the adverse event was the cause of the defect and fluoride
267 was not the cause. On the other hand, if an adverse condition could not be matched to
268 a supposedly fluorosis defect, one could not be sure that fluoride was the cause. In fact,
269 the definite diagnosis of fluorosis is not possible by this method and the only exact way
270 to diagnose dental fluorosis would be by microscopic or chemical analysis. However,
271 although the ECEL method cannot prove that fluoride is or is not the cause of defect, it
272 can help in differentiate between genuine fluorosis and fluorosis-resembling defects. In
273 the case of this study, past childhood data obtained by the ECEL method helped to find
274 fluorosis-resembling defects in 10.2%, 8.5%, and 6.7% of subjects from high, optimal,
275 and low fluoride areas respectively.

276

277 It was decided from the beginning of the study that children with large restorations on
278 their permanent anterior teeth should be excluded as it was not possible to assess the
279 previous existence of defects on them. This was while the authors were aware of the
280 bias this could make. The large restorations could be due to fluorosis, non-fluoride
281 related DDEs, dental caries, trauma or other reasons. One could rule out dental caries
282 and trauma easily by questioning the parents. However, there was no means of
283 distinguishing between fluorosis and any other types of defects. Fortunately, in this
284 study, only one child was excluded based on having large restorations. This one subject

285 could not affect the results of this study. However, this should be addressed in future
286 studies.

287

288 The diagnosis of fluorosis is more complicated than acknowledged. On the other hand,
289 there has always been considerable controversy regarding the use of fluoride for dental
290 caries prevention. One of the main issues, which opponents of fluoridation raise, is
291 fluorosis. Therefore, finding a reliable way for more accurate diagnosis of genuine
292 fluorosis is vital. Existing fluorosis indices could lead to misjudgment about using
293 fluoride. However, the ECEL method to record health-related life events is a promising
294 method to help differentiate between genuine fluorosis and fluorosis-resembling defects.

295

296 **Conclusion:**

297 Fluorosis indices, if used alone, could result in misdiagnosis of dental fluorosis and
298 misguide health policymakers in their decision about public health measure related to
299 use of fluoride. Information about adverse health-related conditions linked to DDEs at
300 specific positions on teeth could help to differentiate between genuine fluorosis and
301 fluorosis-resembling defects.

302

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308

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372

374 Table1. The comparison of fluorosis scores according to Dean's Index by four calibrated
 375 dentists

	Test	Examiner 2	Examiner 3	Examiner 4
Examiner 1	Kappa value	0.16	0.16	0.07
	McNemar p-value	<0.001	<0.001	<0.001
	Correlation coefficient	0.528	0.454	0.458
Examiner 2	Kappa value		0.34	0.37
	McNemar p-value		0.125	<0.001
	Correlation coefficient		0.584	0.575
Examiner 3	Kappa value			0.29
	McNemar p-value			0.003
	Correlation coefficient			0.614

376

378 Table 2.The comparison of fluorosis scores according to TF Index by four calibrated
379 dentists

	Test	Examiner 2	Examiner 3	Examiner 4
Examiner 1	Kappa value	0.06	0.21	0.34
	McNemar p-value	<0.001	<0.001	<0.001
	Correlation coefficient	0.381	0.465	0.498
Examiner 2	Kappa value		0.27	0.18
	McNemar p-value		<0.001	<0.001
	Correlation coefficient		0.472	0.503
Examiner 3	Kappa value			0.36
	McNemar p-value			<0.001
	Correlation coefficient			0.539

380

381

382 Table 3: Comparison of all defects diagnosed as fluorosis, genuine fluorosis, and
 383 fluorosis-resembling defects among the three areas

	All children diagnosed with fluorosis*	Children with genuine fluorosis	Children with fluorosis resembling defects	
High (N= 88)	51 (58.0%)	42 (47.7%)	9 (10.2%)	
Optimal (N=189)	55 (29.1%)	39 (20.6%)	16 (8.5%)	
Low (N= 90)	9 (10.0%)	3 (3.3%)	6 (6.7%)	
Level of fluoride in area	OR optimal/high (95% CI)	0.303** (0.179-0.514)	0.292** (0.168-0.506)	0.797 (0.337- 1.886)
	OR low/high (95% CI)	0.080** (0.036-0.180)	0.037** (0.011-0.127)	0.625 (0.231- 1.837)
Boys (N= 196)	67 (34.2%)	52 (26.5%)	15 (7.7%)	
Girls (N= 171)	48 (28.1%)	32 (18.7%)	16 (9.4%)	
Sex	OR girls/boys (95% CI)	0.744 (0.462-1.201)	0.619 (0.362-1.056)	1.251 (0.597-2.620)

384 * All children that were diagnosed as having fluorosis by three out of four calibrated
 385 dentists using both Dean and TF Indices.

386 ** Significant at 0.001 level.

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