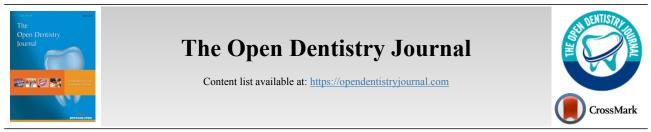
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RESEARCH ARTICLE

The Prevalence and Etiology of Molar-Incisor Hypomineralization among Female School Children

Ghofran S. Sulaimani¹, Lama O. Hejazi¹ and Hanadi S. Lingawi^{1,*}

¹Department of Preventive Dentistry, College of Dentistry, Umm Al-Qura University, Makkah, Saudi Arabia

Abstract:

Background:

Molar-Incisor Hypomineralization (MIH) is a developmental defect affecting teeth. The prevalence of MIH and its clinical implications form a challenge for patients and dentists.

Objective:

The purpose of this study was to determine the prevalence of MIH among female school children in Makkah, Saudi Arabia.

Methods:

A cross-sectional study was conducted including 7- to 9-year-old school girls from the public and private sectors. Invitations to participate in the study were sent to the parents of 1,000 randomly selected children, along with questionnaire and consent form. All included children were examined by two dentists who had been previously calibrated in the assessment of MIH (K=0.9), which was diagnosed according to the European Academy of Paediatric Dentistry 2003 criteria. Data analysis was done using SPSS V.22.0 and various statistical tests were conducted to determine significant differences at P<0.05.

Results:

A total of 711 children participated in this study, with a response rate of 82.2%. The prevalence rates of MIH and Molar Hypomineralization (MH) were 20.96% and 26.58%, respectively. The most common form of MIH was the mild form (87.24%), while 18 children (12.08%) were diagnosed with the moderate form. A single child (0.67%) was diagnosed with the severe form. No statistically significant associations were found between MIH/MH and prenatal, perinatal or postnatal factors.

Conclusion:

The prevalence of MIH in school girls in Makkah was 20.96%. Despite its high prevalence, the severity of MIH was relatively mild. The etiology of MIH remains unclear and warrants further investigation in future studies.

Keywords: Molar-incisor hypomineralization, School children, Enamel hypoplasia, Dental anomalies, Prevalence, Prevalence rate.

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1. INTRODUCTION

Since the 1970s, many cases of developmental defects localized to the First Permanent Molar(s) (FPMs) and incisor(s) have been reported by dentists [1]. Clinical descriptions of these reported cases included opaque enamel with colour ranging from white to yellowish brown and areas with no mineralization or hypomineralization of enamel that varied according to condition severity. Enamel breakdown indicates more severe conditions.

In 2001, the term Molar-Incisor Hypomineralization (MIH) was introduced by Weerheijm *et al.* [2]. Molar-Incisor Hypomineralization is defined as a hypomineralization of systemic origin of one to four FPMs that is frequently associated with affected incisors [2]. The lesions in the FPMs are often associated with maxillary incisors and more rarely with mandibular incisors. When only FPMs are affected while the incisors are spared, the condition is called Molar Hypomineralization (MH) [2].

MIH is usually attributed to systemic insults during the mineralization of the FPM and incisor crowns in the first few years of a child's life [2]. The severity of MIH (mild, moderate or severe) may vary between different teeth in the same

^{*} Address correspondence to this author at the Department of Preventive Dentistry, College of Dentistry, Umm Al-Qura University, P.O. Box 14405, Makkah 22915, Saudi Arabia; Tel. +966 505682076; E-mail: hslingawi@uqu.edu.sa

individual [3] and is a challenging condition for dentists as well as patients. It is usually associated with serious complications such as severe discomfort, hypersensitivity, shooting pain, early loss of fillings and post-eruptive breakdown due to the hypomineralized tooth structure. Children affected with MIH are more susceptible to dental caries and have more Decayed, Missing and Filled Teeth (DMFT) compared to other children with different types of enamel defects [4].

MIH has received increasing attention in the literature in recent years, particularly in Europe, Australia and South America. The worldwide prevalence of MIH has been reported to range from 2.3% in Egypt [5] to 40.2% in Brazil [6]. The wide variation in MIH prevalence rates may be attributed not only to the different geographical regions studied but also to the assessment of different age groups and the use of different indices and criteria for the evaluation of hypomineralization; this has precluded valid comparisons across the studies.

In terms of the Middle East region, prior studies have reported the prevalence of MIH in Libya [7], Egypt [5], Jordon [8], Lebanon [9] and Iraq [10, 11]. Very few studies have been conducted on the Arabian Peninsula. Examples include studies conducted in Bahrain [12] and the United Arab Emirates [13]. In addition, two studies have been conducted among school children in Saudi Arabia; these studies reported MIH prevalence rates of 8.6% and 40.7% in the cities of Jeddah [14] and Riyadh [15], respectively. Notably, these two studies utilised different protocols for MIH assessment and included participants with different age ranges. Furthermore, the study conducted in Jeddah utilised a small sample size; thus, caution must be used in generalising its results to the nation as a whole.

While the number of studies aimed at elucidating the etiological factors of MIH has increased in recent years, the results have been inconclusive [16 - 18]. Nevertheless, it is widely accepted that the etiology of MIH is multifactorial and includes both genetic and environmental factors [17, 18]. Environmental postnatal factors include fevers, chickenpox, tonsillitis, asthma, repeated ear infections, allergies and the repeated use of antibiotics during the first 4 years of life. Prenatal factors include illnesses during pregnancy, birth complications and premature birth.

The dental treatment of MIH/MH cases has proven to be a challenge for both patients and dentists. The porosity of dental enamel leads to pulpal inflammation, and the increased number of nerve endings in dentinal tubules results in sensitivity and pain. Additionally, there are often difficulties in achieving adequate anesthesia. As a result, children with MIH/MH usually experience dental fear and anxiety. Therefore, it is important to determine the prevalence of MIH in a given population, as this will facilitate the development and implementation of policies for its prevention and management. The aim of this study was to determine the prevalence of MIH among female school children in Makkah, Saudi Arabia, and to investigate the possible associated etiological factors.

2. MATERIALS AND METHODS

2.1. Study Design and Ethical Considerations

The study protocol was approved by the Institutional

Review Board (IRB) of the College of Dentistry, Umm Al-Qura University (IRB no. 87-18). This study utilised a crosssectional design and was conducted among 10 primary private and public schools for girls in Makkah, Saudi Arabia. Schools from both private and public sectors were selected to ensure that all socio-economic statuses are included in the study. The schools were selected through stratified random selection to cover different districts. The 10 schools were invited to participate in the study *via* official letters from the Regional Director of the School Health Department in Makkah.

After a thorough review of the literature, a questionnaire was carefully constructed and designed by the authors to document possible etiological factors associated with MIH/MH. It covered four domains: (1) demographic data, (2) maternal health and medications administered during pregnancy, (3) delivery history and (4) the child's medical history during her first 3 years of life (Fig. 1).

Prior to questionnaire distribution, a pilot study to test the precision of the survey questions was conducted with 10 participants. Based on their responses, 4 survey questions were modified, and the format was edited for clarity and comprehensibility. A second pilot study was conducted with another 10 participants to confirm the feasibility, validity and reliability of the questionnaire. A total of 1,000 parental consent forms and questionnaires were distributed to the participating schools. The number of schools and sample size was adjusted to the time limitations of the study, the limited number of clinical examiners (2 dentists) and the research budget.

The sample inclusion criteria comprised the following: (1) females aged 7-9 years, (2) born and brought up in Saudi Arabia and (3) children having at least one erupted or partially erupted FPM. Participants were excluded if they had (1) generalized opacities present on all teeth rather than those limited to the FPMs and permanent incisors such as in case of amelogenesis imperfecta, (2) cases of fluorosis, which is usually diffuse and generalized affecting other than the target teeth, (3) defects in the permanent dentition due to prior dental trauma/ infection in the primary dentition.

2.2. Clinical Examination and Diagnosis

The clinical examinations were performed by two general dentists who had been previously trained and calibrated (using photographs of clinical cases) in the identification of MIH according to the European Academy of Paediatric Dentistry (EAPD) 2003 criteria [19]. MIH severity was determined according to the Wetzel and Reckel criteria (Table 1) [3], and inter-examiner reliability was determined using Cohen's kappa coefficient prior to study commencement; a high level of reliability (K = 0.9) was documented. Children were examined at their respective school premises by one of the two trained examiners. The intraoral examinations were carried out within a classroom setting and conducted with a headlight and disposable dental examination kit that contained the following items: a dental mirror, tweezers, dental explorer and multiple cotton rolls.

UQU DENT School initial	Serial Code School initial
Questionnaire	
Molar-Incisor Hypomineralization	III- History of child delivery
Mona mesor ryponinier and doin	Did the child suffer from any of these conditions during birth?
	- Early delivery 🛛 Yes 🖓 No
	- Low birth weight
Dear parents,	- Congenital condition 🗌 Yes 🗌 No
This questionnaire aims to conduct a research about the prevalence and etiology of Molar-Incisor Hypomineralization	- Ear infection 🛛 Yes 🖓 No
in space school children in Makkah, Saudi Arabia. We value you time and very much appreciate your participation	- Pneumonia 🗌 Yes 🗌 No - Fever 🗌 Yes 🗌 No
in this study.	- Other Condition
Please answer the following questions by ticking the appropriate box 1. Child age 7 years 9 years 2. Place of birth 3 suid Anabia 0 Others	If yes, please specify
3. Nationality 🛛 Saudi 🔅 Non-Saudi	I- Did the child suffer from any of the following conditions during his first 3 years of life? Adenoiditis Yes N No
1- Did the mother experience any of the following conditions during pregnancy?	- Asthma Ves No
1- Did the mouter experience any of the following conditions during pregnancy?	- Chicken Pox - Yes No - Fever Yes No
- Hypotension Anemia (low blood pressure) 🛛 Yes 🖓 No	- Gastrointestinal diseases
- Gastrointestinal diseases during third trimester 🛛 Yes 🗆 No	- Inflammatory skin diseases 🛛 Yes 🖓 No
Gestational Diabetes Yes No Psychological stresses Yes No	- Jaundice Otitis media
	- Measles
	- Repeated tonsillitis
2- Did the mother experience any other conditions during pregnancy?	- Urinary tract infections 🛛 Yes 🗋 No
🗆 Yes 🛛 No	- Other Conditions 🛛 Yes 🖓 No
If yes, please specify	If yes, please specify
- Troubles required labor induction	2- Did the child repeatedly use any of the following?
- Hypoxia 🛛 Yes 🖓 No	
- Any other condition 🛛 Yes 🖓 No	- Antibiotics 🛛 Yes 🖓 No - Fluoride 🖓 Yes 🖓 No
	- Other medications
If yes, please specify	If yes, please specify
	· · · · · · · · · · · · · · · · · · ·
4. Was the mother on any medications during pregnancy	Thank you,
□ Yes □ No	
If yes, please specify	
	2
1	

Fig. (1). Sample of the questionnaire used in the study.

Table 1. Wetzel and Reckel scale of MIH severity.

Degree 1 (mild)	Degree 2 (moderate)	Degree 3 (severe)
Isolated white and	Hypomineralized	Large-scale mineral
cream to yellowish-	yellowish-brown	deficiency with distinct
brown discolorations	enamel on the	yellowish-brown
on the occlusal	majority of the cusps,	discolorations and
surface and coronal	but with only a slight	defects in crown
third of the crown.	loss of substance.	morphology resulting
		from extensive loss of
		enamel.

2.3. Statistical Analysis

Data were entered into a Microsoft Excel spreadsheet and analysed using the Statistical Package of Social Sciences program (SPSS 22.0, Inc., Chicago, USA). Descriptive statistics were used to describe the data in which frequency and percentage for categorical variables were determined. Patient characteristics were presented descriptively, and the frequencies of potential etiological factors were compared between children with MH and MIH using Pearson's chisquare test. A critical probability value (P-value) of <0.05 was used as the cut-off level for statistical significance.

3. RESULTS

Parents of 865 children provided written informed consent and completed/returned the questionnaires (response rate = 86.5%); however, not all of the children were present on the day of the clinical examinations, resulting in a final sample size of 711. The mean age of the recruited children was 8.1 ± 0.73 ; 536 (75.39%) were Saudis and 175 (24.61%) were non-Saudis.

A total of 149 children (20.96%) were diagnosed with MIH, while 189 (26.58%) were diagnosed with MH. Among Saudi children, the prevalence rates of MIH and MH were 21.83% (n = 117) and 27.24% (n = 146), respectively (Fig. 2). Lower prevalence rates of MIH (18.29%; n = 32) and MH (24.57%; n = 43) were observed among non-Saudi children (Fig. 3).

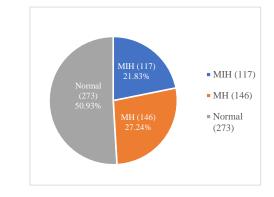


Fig (2). The prevalence of MIH and MH reported among *Saudi* female school children (MIH: molar incisor hypomineralization; MH: molar hypomineralization).

In terms of MIH severity, 130 children (87.25%) were diagnosed with mild MIH, while 18 children (12.08%) were diagnosed with moderate MIH. A single child (0.67%) was diagnosed with severe MIH (Table 2). No severe cases of MH were documented; 187 (98.94%) and 2 children (1.06%) were diagnosed with mild and moderate MH, respectively.

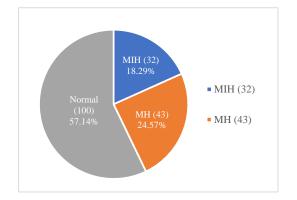


Fig. (3). The prevalence of MIH and MH reported among *non-Saudi* female school children (MIH: molar incisor hypomineralization; MH: molar hypomineralization).

No significant associations were found between MIH/MH and pre-natal factors (*i.e.*, birth complications, illnesses during pregnancy or birth complications) or illnesses affecting the child during her first 3 years of life (*e.g.* fever, jaundice, otitis media, or frequent tonsillitis) (Table **3**). Frequent antibiotic use was more commonly reported for children with MH (46.5%) compared to children with MIH (42.9%); nevertheless, this did not reach statistical significance (*p* value = 0.074) (Table **3**).

4. DISCUSSION

Prevalence studies play a major role when strategies and policies need to be developed to prevent/treat certain conditions. This study represented the first MIH prevalence and etiology study conducted in the city of Makkah and the third of its type in Saudi Arabia [14, 15].

Both the response (86.5%) and final participation rates (82.2%) were good; therefore, the recruited sample was considered to be representative of the target population. The age range chosen in this study (7-9 years old) was considered suitable for MIH assessment, as the four FPMs would have erupted by this age in most children. A similar age range has been used in other studies, such as in Zawaideh et al.'s study of MIH in Jordanian children, in Petrou et al.'s study of MIH prevalence among German school children, and in Mittal et al. [8, 20, 21]. This age range criterion also minimized the risk that the areas of enamel hypomineralization would be masked by subsequent dental caries or restorations. The selection criteria of this study excluded children with generalized opacities present on all teeth, fluorosis or defects in the permanent dentition due to prior dental trauma/ infection in the primary dentition, as these conditions represent a grey area in the diagnoses, where MIH/ MH might be masked or misdiagnosed with other conditions. For instance, diffuse

opacity may be caused by the intake of a high concentration of fluoride and should not, according to Weerheijm *et al.* [19], be classified as MIH.

 Table 2. Prevalence of MIH severity degrees in the studied sample.

MIH severity	Frequency (n)	Percentage (%)	X^2	p value	
Mild	130	87.25	748.203	0.000	
Moderate	18	12.08			
Severe	1	0.67			
TOTAL	149	100.00			
MIH: molar incisor hypomineralization.					

In an effort to ensure the consistency and reliability of the clinical assessments, all examinations were carried out by two general dentists who had been previously trained and calibrated.

The results of our study showed that 338 children (47.5%) of all examined children had hypomineralization of at least one examined tooth. The prevalence of MIH in our study was 20.96%, which was greater than that reported by Allazzam et al. (8.6%) in a study conducted in Jeddah, which is located on Saudi Arabia's west coast [14]. This difference may have been attributed to their use of a convenience sample obtained at a single university teaching hospital, as well as a small sample size of 267 children. Indeed, Elfrink et al. [22] suggested that a minimum random sample size of 300 participants would be required for an accurate estimation of MIH prevalence. Al-Hammad et al. [15] reported a much higher prevalence of MIH (40.7%) in the city of Riyadh, located in the middle of Saudi Arabia. While this may be due to true regional variation, the effect of differences in study methodology must also be acknowledged; this emphasises the importance of conducting a national survey for the accurate estimation of MIH prevalence in Saudi Arabia.

Based on the results of the present study, MIH in Saudi Arabia was considered to be moderately prevalent relative to other countries around the world, such as Brazil (40.2%) [6], Denmark (37.5%) [23], Finland (25%) [24] and Egypt (3.2%) [5]. However, studies conducted in the Middle East region have consistently reported lower prevalence rates; examples include Libya (2.9%) [7], Jordon (17.6%) [8], Bahrain (17.6%) [12] and Iraq (18.6%) [11, 12]. The only country in the Gulf region with a higher reported MIH prevalence than the present study, was the United Arab Emirates (34.4%) [13]; this may be attributed to the differences in inclusion criteria between both studies and the use of a smaller sample size (less than half that of the present study).

This study is considered one of the very few studies that studied the prevalence of both MIH and MH in the same population. Most of the studies investigated the prevalence of MIH. The results of our study showed a higher prevalence of MIH (20.96%) than MH (26.58%) in the studied sample, which goes in agreement with the results reported by *Hussien et al.* [13]. Their results reported that the prevalence of MH was higher than MIH (65.6% and 34.4%), respectively.

Condition	MIH		MH		Not Affected		<i>p</i> Value
	Frequency (n)	Percent (%)	Frequency (n)	Percent (%)	Frequency (n)	Percent (%)	
Birth complications	9	6.04	14	7.41	25	6.70	0.574
Repeated attacks of tonsillitis	41	27.52	64	33.86	120	32.17	0.769
Otitis media	18	12.08	24	12.70	49	13.14	0.320
Skin allergy	22	14.67	38	20.11	79	21.18	0.211
Fever	70	46.98	101	53.44	189	50.67	0.303
Asthma	20	13.42	44	23.28	87	23.32	0.283
Jaundice	12	8.05	18	9.52	35	9.38	0.323
Repeated antibiotic intake	64	42.90	88	46.56	189	50.70	0.074
Total	149	100	189	100	373	100	

Table 3. The association between possible etiological factors and the prevalence of MIH/ MH.

MH: Molar Hypomineralization; MIH: Molar Incisor Hypomineralization.

The finding that the majority of the MIH and MH cases were of the mild form is in agreement with the results of other studies conducted among the same age group [8, 20]. However, this might be age-related. It has been well-established that hypomineralized teeth are weaker and more susceptible to post-eruptive enamel breakdown and dental caries. Hence, an increase in hypomineralization severity has been found to be associated with an increase in age [25, 25].

It is notable that the prevalence rates of caries and fluorosis are high in the western region of Saudi Arabia [26, 27], as both conditions hinder the assessment of MIH. In order to address this issue, we targeted a younger age group (7-9 years old) in the present study; this allowed for the assessment of permanent teeth soon after their eruption, thus minimizing the probability of existing caries and restorations. Similar to previous studies [8, 13], we observed a higher prevalence of MH (26.58%) than MIH (20.96%). This result may have been attributed to the earlier completion of molar formation and calcification compared to incisors [28 - 30].

In terms of the investigation of possible etiological factors for hypomineralization, 663 children did not have birth complications nor hypomineralization; 25 had birth complications but no hypomineralization; and 23 had birth complications and hypomineralization. Almost 47.9% of our sample population who had birth complications also had hypomineralization. It is justifiable that MH had higher association with birth complications (14 children) than MIH (9 children). This is because the first permanent molars begin mineralizing around birth, while the mineralization of the first permanent incisors occurs during the first year of life. However, the results of the present study indicated no statistically significant association between prenatal/perinatal factors and MIH/MH [31, 32]. Specifically, two prior studies have also reported the lack of a significant association between birth complications and MIH/MH [18, 32], supporting the results of our study. On the other hand, childhood illnesses during the first 3 years of life have been reported to be more common among children with MIH or MH [14, 32]. For instance, fever and asthma in the first 3 years of life have been reported to be associated with MIH [33 - 35]. Nevertheless, no significant association was found between MIH/MH and either fever or asthma in the present study.

To date, the relationship between MIH/MH and repeated use of antibiotics in the first 3 years of life has been controversial. There is also uncertainty about whether the potential effects on hypomineralization are limited to amoxicillin antibiotics or whether all antibiotic types may be implicated. In the present study, the association between hypomineralization and frequent antibiotic use during the first 3 years of life did not reach statistical significance. This was supported by a previous study conducted in Spain, which did not find a significant relationship between antibiotic consumption and MIH/MH [26]. On the other hand, a study conducted in the United Kingdom reported a significant association between MIH and the frequent use of amoxicillin between the ages of 3 and 3.9 years [18]. This result has remained controversial, as other evidence has suggested that the type of antibiotic is not as important as the timing of administration (i.e. during the first 3 years of life) [10]. At any rate, it is difficult to ascertain whether hypomineralization is associated with the prescription of antibiotics to treat an existing infection or the effects of the infection itself.

The main strength of the present study was the large sample size employed, which was the largest to date among studies conducted in the Arabian Gulf countries. A possible limitation was that the questionnaires were self-completed by the parents, and no interviewer was utilized to obtain the responses. This may have compromised the reliability of the answers. Furthermore, the recruited sample was limited to females; a future study utilizing an identical protocol in boys' schools is currently being planned. This will allow the determination of sex differences in MIH/MH prevalence and their potential interaction with other risk factors.

CONCLUSION

In conclusion, the prevalence rates of MIH and MH were determined to be 20.96% and 26.58%, respectively, in 7- to 9-year-old females in Makkah, Saudi Arabia. Both hypomineralization types were mainly mild in severity. There was no statistically significant association between any of the investigated etiological factors and MIH/MH. Nevertheless, additional studies are required to further elucidate the association of prenatal, perinatal and postnatal effects on hypomineralization.

ETHICS APPROVAL AND CONSENT TO PARTI-CIPATE

The study received ethical approval from the Institutional Review Board (IRB) at the College of Dentistry, Umm Al-Qura University, Saudi Arabia (IRB No. 87-18).

HUMAN AND ANIMAL RIGHTS

No animals were used in this research. All human research procedures followed were in accordance with the ethical standards of the committee responsible for human experimentation (institutional and national), and with the Helsinki Declaration of 1975, as revised in 2013.

CONSENT FOR PUBLICATION

Parents of 865 children provided written informed consent.

AVAILABILITY OF DATA AND MATERIALS

All data that support the results of this study are restricted by the IRB, College of Dentistry, Umm Al-Qura University in order to protect patient privacy. Data can be made available upon reasonable request from the IRB secretary (irb.uqudent@uqu.edu.sa) only for researchers who meet the criteria for access to confidential data.

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None.

CONFLICT OF INTEREST

The authors declare no conflict of interest, financial or otherwise.

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Declared none.

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