

Prevalence and Patterns of Non-syndromic Hypodontia in Permanent Dentition among the Nepalese Population: A Radiographic Study and Literature Review



Shantanu Dixit^{1,*}, Dinesh Rokaya^{2,3} and Muhamad Nizam Muhamad Subra⁴

¹Department of Oral Medicine and Radiology, Faculty of Dental Sciences, Shree Guru Gobind Singh Tricentenary (SGT) University, Gurugram, India

²Clinical Sciences Department, College of Dentistry, Ajman University, Ajman, United Arab Emirates

³Center of Medical and Bio Allied Health Sciences Research, Ajman University, Ajman, United Arab Emirates

⁴Department of Orthodontics, Faculty of Dentistry, Universiti Teknologi MARA, Selangor, Malaysia

Abstract:

Introduction: Odontogenesis is a complex process involving interactions between odontogenic and ectomesenchymal cells, mediated by growth factors and signaling pathways. Disruptions in these pathways lead to various dental anomalies. This study aimed to evaluate the prevalence and patterns of non-syndromic hypodontia among the Nepalese population.

Methods: A total of 5,075 panoramic radiographs taken between 2012 and 2020 were retrospectively reviewed. The mean age of the study subjects was 20.17 ± 5.51 years (range: 8 to 40 age), comprising 1,805 males (35.56%) and 3,270 females (64.43%). Hypodontia was diagnosed based on the absence of crown calcification of permanent teeth. Prevalence, distribution, and symmetry of hypodontia were assessed, with categorical associations analyzed using the Chi-square test ($p < 0.05$).

Results: The study analyzed 5,075 panoramic radiographs, consisting of 1,805 males (35.56%) and 3,270 females (64.43%). Hypodontia was identified in 400 subjects, 200 males (50%) and 200 females (50%), representing 7.88% of the population. A total of 655 missing teeth were recorded, 45.8% in females and 54.2% in males. Despite an equal number of affected males and females, a statistically significant association was found between gender and overall hypodontia prevalence in the sample ($p < 0.05$), with males showing a higher likelihood of being affected. Maxillary lateral incisors were the most affected (47.33%), followed by mandibular second premolars (19.08%) and maxillary first premolars (17.56%). The maxillary arch had a significantly higher prevalence of hypodontia than the mandibular arch ($p < 0.05$). Unilateral hypodontia was more prevalent than bilateral hypodontia ($p < 0.05$). The most common associated dental anomalies were rotated adjacent teeth and retained deciduous teeth, but they were not statistically significant ($p > 0.05$).

Conclusion: The prevalence of hypodontia among the Nepalese population was 7.8%. The findings of this study align with global trends, particularly the higher frequency of hypodontia in the maxillary lateral incisors.

Keywords: Hypodontia, Tooth agenesis, Prevalence, Dental anomalies, Nepal, Odontogenesis.

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*Address correspondence to this author at the Department of Oral Medicine and Radiology, Faculty of Dental Sciences, Shree Guru Gobind Singh Tricentenary (SGT) University, Gurugram, India; E-mail: drshantanu86@gmail.com

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1. INTRODUCTION

Odontogenesis, or tooth development, is an intricate process encompassing a series of sequential and reciprocal interactions between odontogenic and ecto-mesenchymal cells derived from the stomadeal epithelium and cranial neural crest cells, respectively [1]. These interactions are intermediated by growth and transcription factors, signal receptors, and numerous soluble morphogens [2]. The developing tooth endures a multitude of morphological, histological, and biochemical transformations to attain physiological and functional maturity [3].

The impediment of typically impeccable phenomena of odontogenesis leads to the emergence of numerous dental anomalies [4]. Tooth agenesis (TA) is one of the prevalent developmental anomalies impacting human dentition. It is defined as the developmental absence of one or more teeth from the 'normal' set of human dentition, typically excluding permanent third molars. The number of developmentally absent teeth further classifies TA into three distinct types: hypodontia (absence of 1-5 teeth), oligodontia (absence of more than 6 teeth), and anodontia (agenesis of all teeth) [5].

Hypodontia, also referred to as 'selective tooth agenesis' or 'congenital tooth absence', predominantly affects permanent dentition rather than deciduous dentition [6]. The lack of a tooth's eruption in the oral cavity, in conjunction with the absence of radiographic evidence supporting its existence within the anticipated period, suggests a diagnosis of hypodontia [7]. This anomaly can manifest itself as a part of a genetic syndrome (like anhidrotic ectodermal dysplasia, Down syndrome, Ehlers-Danlos syndrome, Van der Woude syndrome, etc.) or as an isolated trait of the non-syndromic familial form [5].

Brook's unifying etiological model for anomalies in tooth number and size posits that hypodontia arises from complex interactions among genetic, epigenetic, and environmental influences throughout the process of odontogenesis [8]. Various environmental variables, including radiation, chemotherapeutic agents, and infections, can impede this process. However, the genetic element prevails over the environmental variables in the etiology of hypodontia. Molecular investigations substantiate this assertion by linking it to mutations in genes active during the initial stages of odontogenesis, such as paired box 9 (PAX9), muscle segment homeobox 1 (MSX1), axis inhibitor 2 (AXIN2), and ectodysplasin A (EDA) [9]. Furthermore, epigenetic alterations, such as DNA methylation and histone modification, influence a network of interrelated signaling pathways that encompass odontogenesis, subsequently leading to hypodontia [10].

Non-syndromic hypodontia is reported to be relatively more prevalent than its syndromic counterpart [11]. Khalaf *et al.* [12] conducted a meta-analysis and estimated the overall prevalence rate of this anomaly to be 6.4%, with a statistically significant difference across various continents. For this anomaly, Africa reported the highest prevalence rate of 13.4%. Europe recorded a prevalence rate of 7%, followed closely by Asia and Australia, both at 6.3%. In contrast, North America exhibited a lower rate of 5.0%, while Latin America and the Caribbean had an even lesser prevalence at 4.4%. Furthermore, this analysis showed that

hypodontia was more widespread in females, with a total female: male ratio of 1.22:1. In addition, it was predominantly observed in mandibular second premolars, followed by maxillary lateral incisors and maxillary second premolars. A meta-analysis conducted by Polder *et al.* [13] further highlighted that unilateral hypodontia is more prevalent than its bilateral counterpart.

The prevalence of hypodontia is known to vary based on factors, such as patients' ethnicity, sampling techniques, diagnostic criteria, chronological age, and sample size [11]. Hence, the present study evaluated the prevalence and pattern of non-syndromic hypodontia of permanent teeth in a subgroup of the Nepalese population. Additionally, a targeted literature review was carried out to compare the prevalence patterns, diagnostic methods, and demographic variations observed in different populations worldwide. Furthermore, this comparative analysis sought to provide a broader understanding of hypodontia trends and their epidemiological significance.

This study aimed to address a critical gap in the existing body of literature, with the anticipation that its findings would provide a robust foundation to inform and guide future research endeavors.

2. MATERIALS AND METHODS

2.1. Study Design

This is a retrospective cross-sectional study that was conducted at the University Hospital of Kathmandu University School of Medical Sciences, Dhulikhel (Kavre, Nepal), utilizing panoramic radiographs. The study adhered to the principles of the Declaration of Helsinki and received approval from the Institutional Review Board (IRB Protocol approval no. 61/19).

2.2. Study Population

This study employed a convenience sampling approach to select panoramic radiographs from the digital archives of the Department of Oral Radiology. Given the retrospective nature of the study, this sampling method was deemed appropriate as it allowed for efficient data collection from a pre-existing institutional dataset spanning eight years (2012-2020). Convenience sampling, while non-random, provided access to a substantial and diverse sample, facilitating a comprehensive evaluation of the prevalence and patterns of hypodontia within the study population.

The selection of radiographs was based on strict inclusion and exclusion criteria to ensure the validity and reliability of the findings. The inclusion criteria comprised panoramic radiographs of individuals aged 8 to 40 years with permanent dentition and no history of systemic syndromes or craniofacial abnormalities. To maintain diagnostic accuracy, only high-quality radiographs free from distortion or artifacts were considered. Conversely, the exclusion criteria encompassed radiographs that exhibited missing teeth due to trauma, extractions, or orthodontic treatment, as these factors could confound the assessment of congenital tooth absence. Additionally, radiographs of individuals diagnosed with syndromic conditions known to affect dental development were excluded to maintain the study's focus on non-syndromic hypodontia.

Following the application of these criteria, a total of 5,075 panoramic radiographs were included in the study, comprising 1,805 males (35.56%) and 3,270 females (64.43%) aged 8 to 40 (20.17 ± 5.51) years. While convenience sampling does not provide randomization, its application in this study enabled the inclusion of a large dataset over an extended timeframe, thereby reducing potential biases and enhancing the study's statistical robustness.

Hypodontia was diagnosed by the absence of crown calcification of one or more permanent teeth (excluding third molars) on radiographic examination. The teeth evaluated included all permanent maxillary and mandibular teeth from the central incisors to the second molars (*i.e.*, teeth numbered 11-17, 21-27, 31-37, and 41-47 according to the FDI two-digit system). A diagnosis of hypodontia was established when there was no radiographic evidence of crown calcification for a given tooth and no documented history of its extraction, loss due to trauma, or orthodontic removal.

Two independent observers (SD and DR) meticulously evaluated all panoramic radiographs to ensure diagnostic accuracy, with any discrepancies resolved through mutual consensus. For each confirmed case of hypodontia, comprehensive data were recorded, including demographic details, such as age (in years) and gender (male or female). Dental variables encompassed the presence or absence of permanent teeth (excluding third molars), the location of missing teeth (maxillary vs. mandibular arch), the specific type of missing teeth (*e.g.*, lateral incisors, second premolars, or first premolars), and the pattern of occurrence (unilateral or bilateral). Additionally, associated dental anomalies were documented where applicable.

2.3. Statistical Analysis

The data was collected and processed utilizing SPSS software, version 21 (IBM Corp., Armonk, N.Y., USA). Descriptive and inferential statistics were used to analyze the data, with descriptive statistics specifically applied to summarize the study variables. Prevalence was calculated as a percentage, and patterns were categorized by tooth type, arch (maxillary or mandibular), and symmetry (unilateral or bilateral). Categorical variables association was determined by the Chi-square test with a significance threshold of 0.05.

3. RESULTS

The study analyzed 5,075 panoramic radiographs, comprising 1,805 males (35.56%) and 3,270 females (64.43%). Hypodontia was identified in 400 subjects: 200 males (50%) and 200 females (50%). A total of 655 permanent teeth (excluding third molars) were missing, with 45.8% (300 teeth) in females and 54.2% (355 teeth) in males.

Although the number of males and females with hypodontia was equal, the overall sample had an unequal gender distribution. Therefore, a Chi-square test was carried out to assess the association between gender and the prevalence of hypodontia in the total population. The result was statistically significant ($p < 0.05$), indicating that males had a significantly higher likelihood of presenting with hypodontia than females when considering the sample's gender composition (Table 1). No cases of oligodontia or anodontia were observed within the investigated cohort.

Out of a total of 655 missing permanent teeth, the lateral incisors were the most frequently affected, comprising 47.33% of cases. This was followed by the second premolars (19.08%) and the first premolars (17.56%). Hypodontia was less common in canines (12.21%) and rare in central incisors (3.82%), with no incidences reported for molars (Fig. 1).

Table 1. Tooth-wise hypodontia in male and female patients.

Gender	Female					Male						χ^2 Test
Tooth Region	Maxillary		Mandible		Sub-total %	Maxillary		Mandible		Sub-total %	Total	-
	Right Side (%)	Left Side (%)	Right Side (%)	Left Side (%)		Right Side (%)	Left Side (%)	Right Side (%)	Left Side (%)			-
Central Incisors	5 (0.76)	5 (0.76)	0 (0)	0 (0)	10 (1.53)	10 (1.53)	5 (0.76)	0 (0)	0 (0)	15 (2.29)	25 (3.82)	$\chi^2 = 4.62$, $df = 1$, $p = 0.032^*$
Lateral Incisors	90 (13.74)	55 (8.39)	0 (0)	5 (0.76)	150 (22.9)	75 (11.45)	55 (8.39)	5 (0.76)	25 (3.82)	160 (24.43)	310 (47.33)	
Canines	0 (0)	20 (3.05)	10 (1.53)	10 (1.53)	40 (6.11)	25 (3.82)	15 (2.29)	0 (0)	0 (0)	40 (6.11)	80 (12.21)	
First Premolar	20 (3.05)	15 (2.29)	5 (0.76)	10 (1.53)	50 (7.63)	25 (3.82)	25 (3.82)	5 (0.76)	10 (1.53)	65 (9.92)	115 (17.56)	
Second Premolar	5 (0.76)	10 (1.53)	10 (1.53)	25 (3.82)	50 (7.63)	15 (2.29)	20 (3.05)	15 (2.29)	25 (3.82)	75 (11.45)	125 (19.08)	
First Molar	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	
Second Molar	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	
Total	120 (18.32)	105 (16.03)	25 (3.82)	50 (7.63)	300 (45.8)	150 (22.9)	120 (18.32)	25 (3.82)	60 (9.16)	355 (54.2)	655 (100)	

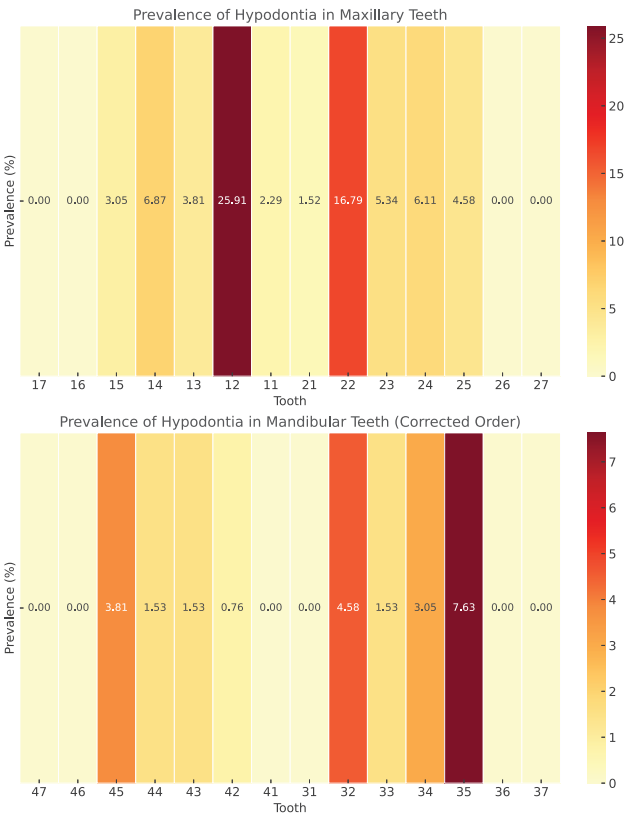


Fig. (1). Tooth-wise prevalence of hypodontia.

Table 2. Unilateral and bilateral patterns of tooth-wise hypodontia in male and female patients.

Tooth Region	Female					Male					Total (%)
	Maxilla (%)		Mandible (%)		Sub-total %	Maxilla (%)		Mandible (%)		Sub-total (%)	-
	Unilateral (%)	Bilateral (%)	Unilateral (%)	Bilateral (%)		Unilateral (%)	Bilateral (%)	Unilateral (%)	Bilateral (%)		
Central Incisors	8 (1.22)	2 (0.30)	0 (0)	0 (0)	10 (1.53)	11 (1.68)	4 (0.61)	0 (0)	0 (0)	15 (2.29)	25 (3.82)
Lateral Incisors	123 (18.78)	22 (3.56)	5 (0.76)	0 (0)	150 (22.9)	88 (13.44)	42 (6.41)	28 (4.27)	2 (0.31)	160 (24.43)	310 (47.32)
Canines	18 (2.75)	2 (0.30)	18 (2.75)	2 (0.31)	40 (6.11)	38 (5.80)	2 (0.31)	0 (0)	0 (0)	40 (6.11)	80 (12.21)
First Premolar	23 (3.51)	12 (1.83)	15 (2.29)	0 (0)	50 (7.63)	36 (5.49)	14 (2.14)	11 (1.68)	4 (0.61)	65 (9.92)	115 (17.56)
Second Premolar	7 (1.07)	8 (1.22)	21 (2.29)	14 (2.14)	50 (7.63)	33 (5.04)	2 (0.31)	34 (5.19)	6 (0.92)	75 (11.45)	125 (19.08)
First Molar	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
Second Molar	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
Total	179 (27.33)	46 (7.02)	59 (9.01)	16 (2.44)	300 (45.8)	206 (31.45)	64 (9.77)	73 (11.14)	12 (1.83)	355 (54.19)	655 (100)

Furthermore, hypodontia exhibited a significantly higher prevalence in the maxillary arch in contrast to the mandibular arch ($p < 0.05$). Unilateral hypodontia was significantly more prevalent than bilateral hypodontia in both genders ($p < 0.05$) (Table 2). Furthermore, the study

identified dental anomalies in individuals with hypodontia, such as rotated adjacent teeth and retained deciduous teeth (Fig. 2). However, these observations were not statistically significant.

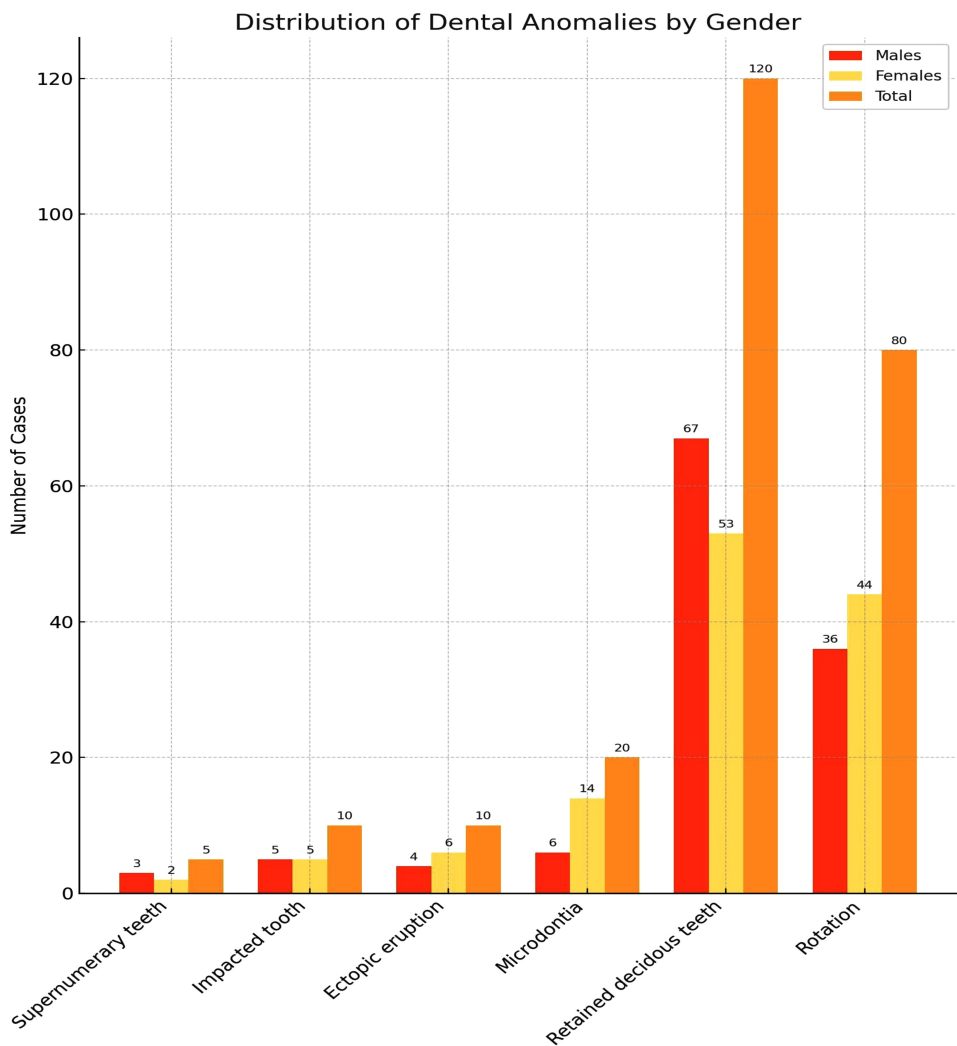


Fig. (2). Graph depicting associated anomalies within the subjects with hypodontia.

4. DISCUSSION

Hypodontia, recognized as the most prevalent cranio-facial anomaly, affects approximately one-fourth of the global population. Its varied genetic and phenotypic manifestations have consistently driven researchers to conduct extensive epidemiological, analytical, and molecular studies aimed at a deeper understanding of this condition [1]. Although numerous studies have explored the prevalence and patterns of hypodontia, there remains a notable lack of data specific to the Nepalese population. To address this gap, the present investigation was undertaken to determine the prevalence of this anomaly within a defined subgroup of the Nepalese population. As part of this study, a targeted literature review was conducted to synthesize existing research on hypodontia. It was performed using keywords, such as “tooth agenesis,” “hypodontia,” “dental agenesis,” “congenitally missing teeth,” and “dental anomalies” across databases, including PubMed, Google Scholar, and Scopus. Studies were selected based on specific inclusion and exclusion criteria: only prevalence-based studies on hypodontia or tooth agenesis (excluding third molars) with a

sample size of 200 or more, published between 2000 and 2024 in English, were considered. Studies focusing on syndromic cases of hypodontia, those with a sample size below 200, non-English publications, case reports, review articles, and studies lacking clear diagnostic criteria were excluded to ensure methodological consistency and comparability.

Studies were further categorized based on the methodology used for hypodontia diagnosis, including radiographic assessments (panoramic or intraoral radiographs), clinical examinations, and journal-reported findings. Additionally, the type of study design (cross-sectional, retrospective, or prospective) was documented to assess the reliability of the findings. The reviewed literature was critically analyzed to compare prevalence rates, gender distribution patterns, and tooth-type involvement across different populations. The insights gained from this literature contribute to a broader understanding of hypodontia trends and provide a comparative framework for evaluating the Nepalese cohort.

A summary of the literature review illustrating the prevalence pattern among different demographics is presented in Table 3 [14-89].

Table 3. Existing data on hypodontia of permanent teeth across diverse populations.

Reference Study	Publication Year	Country	Study Type	Evaluation Method	Age Range (Years)	Total Sample Size	Hypodontia %	Distribution between the Genders %		Most Frequently Reported Hypodontia
								Male	Female	
Present Study	2024	Nepal	Cross-sectional (Retrospective)	Radiographs (panoramic)	8-40	5075	7.8	3.9	3.9	12>22>35>14>24>23>25,32>13,45>15,34>11>43,44,21,33>42
Cavare A <i>et al.</i> [14]	2024	France	Retrospective	Radiographs (panoramic)	9-21	4569	7.9	3.13	4.16	35>45>12>22>45>12>22>42>15>25>41
Katanaki <i>et al.</i> [15]	2024	Greece	Retrospective	Radiographs (panoramic)	9-16	621	5.3	2.41	2.89	35,45>12,22,31,41>33,43>15,25>37,47
Meistere <i>et al.</i> [16]	2024	Latvia	Cross-sectional	Radiographs (panoramic)	11-14	2692	9.3	3.6	5.9	35,45>12,22>15,25
Kanchanasevee <i>et al.</i> [17]	2023	Thailand	Cross-sectional (retrospective)	Radiographs (panoramic)	15.96 ±0.83	1090	9.3	3.39	5.87	45,35>42,32>22,12
Schonberger <i>et al.</i> [18]	2023	Israel	Cross-sectional (retrospective)	Clinical examination, intraoral photographs, dental models, and radiographs (panoramic)	10-25	3000	10.86	4.63	6.23	12,22>35,45>15,25
Vinjolli <i>et al.</i> [19]	2023	Albania	Cross-sectional (retrospective)	Radiographs (panoramic)	15.1±5.5	779	24.39	9.62	14.76	12>22>35>4515>25
Gupta <i>et al.</i> [20]	2022	Nepal	Retrospective	Radiographs (panoramic)	16.42 ± 3.428	601	7.48	Not Specified		12,22>32,42>31,41>35,45>15,25
Eshgian N. <i>et al.</i> [21]	2021	America	Retrospective	Radiographs (panoramic)	Not Specified	1101	2.08	0.72	1.36	14,15,24,25> 12,22,32,42> 34,35,44,45
Yagnam, K <i>et al.</i> [22]	2020	Chile	Cross-sectional	Radiographs (panoramic)	Not Specified	9207	3.02	1.12	1.9	35,45>12,22
Kumar D <i>et al.</i> [23]	2020	India	Cross-sectional	Radiographs (panoramic)	12-28	1100	11.6	5.8	5.8	12,22>31,41>35,45
Fahim And Elabbasy [24]	2020	Egypt	Cross-sectional (retrospective)	Radiographs (panoramic)	14-25	435	10.1	7.6	2.5	12,22>35,45>15,25
Pallikaraki G <i>et al.</i> [25]	2020	Greece	Cross-sectional (retrospective)	Radiographs (panoramic)	7-17	1200	6.6	2.8	3.8	34,44>15,25,14,24>35,45
Bandaru BK <i>et al.</i> [26]	2019	India	Cross-sectional	Clinical examination	3-15	5000	0.8	0.6	0.2	12,22
Aragón HN <i>et al.</i> [27]	2019	Argentina	Cross-sectional (retrospective)	Radiographs (panoramic)	7-11	223	4.48	2.24	2.24	35,45>12,22,15,25>31,41
Haghanifar S., <i>et al.</i> [28]	2019	Iran	Cross-sectional (retrospective)	Radiographs (panoramic)	Not Specified	8018	1.67	0.54	1.13	12,22>15,25
Chandak R <i>et al.</i> [29]	2019	India	Cross-sectional (retrospective)	Radiographs (panoramic)	12-36	1000	1.9	0.9	1	Not specified
Gurbuz O. [30]	2019	Turkey	Cross-sectional	Clinical and Radiographical (panoramic) examination	12-93	2203	3.4	1.27	2.13	Not specified
Aldhorae KA [31]	2019	Yemen	Cross-sectional (retrospective)	Radiographs (panoramic)	9-52	1202	7.48	3.24	4.24	12,22
Fernandez CCA <i>et al.</i> [32]	2018	Brazil	Cross-sectional (retrospective)	Pre-orthodontic records	> 8	1047	7.44	2.67	4.77	24>22>21
Baron C <i>et al.</i> [33]	2018	France	Cross-sectional (retrospective)	Intraoral photographs and radiographs (panoramic)	15.23	551	5.2	1.4	3.8	35,45>15,25>12,22
Hekmatfar S <i>et al.</i> [34]	2018	Iran	Cross-sectional (retrospective)	Radiographs (panoramic)	Not specified	1800	3.72	2.05	1.67	35,45>12,22
Anitha <i>et al.</i> [35]	2018	India	Cross-sectional	Clinical examination	10-70	7018	0.28	Not specified		

(Table 3) contd....

Reference Study	Publication Year	Country	Study Type	Evaluation Method	Age Range (Years)	Total Sample Size	Hypodontia %	Distribution between the Genders %		Most Frequently Reported Hypodontia
								Male	Female	
Roslan AA <i>et al.</i> [36]	2018	Malaysia	Cross-sectional (retrospective)	Dental models, and radiographs (panoramic)	Not specified	370	7	Not specified		32,42>35,45
Chandrika V. <i>et al.</i> [37]	2018	India	Cross-sectional (retrospective)	Dental models, and radiographs (panoramic)	12-30	600	3	0.33	2.67	12,22>15,25>35,45
Sola A. <i>et al.</i> [38]	2018	Spain	Cross-sectional (retrospective)	Radiographs (panoramic)	7-11	2500	3.48	1.8	1.6	35, 45 > 12, 22
Zakaria H <i>et al.</i> [39]	2018	UAE	Cross-sectional (retrospective)	Radiographs (panoramic)	Not specified	2925	1.26	0.65	0.61	Not specified
Ifesanya J.U <i>et al.</i> [40]	2018	Nigeria	Cross-sectional (retrospective)	Radiographs (panoramic)	10-46	216	10.2	Not specified		35,45>12,22>14,15 >32,42
Goutham B <i>et al.</i> [41]	2017	India	Cross-sectional (retrospective)	Radiographs (panoramic)	18-62	1080	4.07	2.22	1.85	Not specified
Almaz M.E. <i>et al.</i> [42]	2017	Turkey	Cross-sectional	Clinical and radiographical (panoramic) examination	1-15	9173	0.52	0.21	0.31	35,45
Laganà G. <i>et al.</i> [43]	2017	Italy	Cross-sectional (retrospective)	Radiographs (panoramic)	8-12	4706	6.69	3.35	3.34	45
Gracco ALT. <i>et al.</i> [44]	2017	Italy	Cross-sectional (retrospective)	Radiographs (panoramic)	9 - 16	4006	9	4.2	5.1	35,45 > 12, 22> 15,25 > 14,24
Badrov J. <i>et al.</i> [45]	2017	Croatia	Cross-sectional (retrospective)	Pre-orthodontic records	6-15	4430	7.8	3.4	4.4	34,45>12,22>15,25
Al-Sheraydah NA <i>et al.</i> [46]	2017	Iraq	Cross-sectional	Clinical examination	15	1000	4.6	1.3	3.3	Not specified
Abdulkareem GB. [47]	2016	Sudan	Cross-sectional (retrospective)	Radiographs (panoramic)	18-45	1225	8	1.55	6.45	35,45>15,25>12,22 >32,42
Kumar A [48]	2016	India	Cross-sectional	Clinical and radiographical (panoramic) examination	14-60	1500	12.8	6.6	6.2	Not specified
Saberi EA <i>et al.</i> [49]	2016	Iran	Cross-sectional (retrospective)	Radiographs (panoramic)	Not specified	1172	1.1	0.59	0.51	12,22>35,45>32,42
Dang HQ <i>et al.</i> [50]	2016	Australia	Cross-sectional (prospective)	Radiographs (panoramic)	6-18	1050	4.28	2	2.28	35,45>12,22>15,25 >37,47>32,42>34,44
Hashim HA <i>et al.</i> [51]	2016	Qatar	Cross-sectional (retrospective)	Pre-orthodontic records	11-36	1000	7.8	2.4	5.4	12,22>34,45>15,25 >32,42
Yassin SM [52]	2016	Saudi Arabia	Cross-sectional (retrospective)	Clinical and Radiographical (panoramic) examination	Not specified	1252	9.66	4.23	5.43	35,45>12,22
Yamunadevi A <i>et al.</i> [53]	2015	India	Cross-sectional	Clinical examination	17-21	244	2	0	2	12,22
Abu-Hussein M <i>et al.</i> [54]	2015	Israel	Cross-sectional (retrospective)	Radiographs (panoramic; intraoral periapical)	12-39	2200	2.59	1.09	1.50	35,45>12,22>15,25 >31,41
Vani N.V. <i>et al.</i> [55]	2015	Saudi Arabia	Cross-sectional (retrospective)	Radiographs (panoramic)	18-40	1000	5.2	2.2	3	12,22>35,45,34,44 >14,24,15,25
Tantanaporn KW. [56]	2015	Thailand	Cross-sectional (prospective)	Clinical and radiographical (Panoramic) examination	13-30	638	13.16	2.82	10.34	31,32,41,42>34,35,44,45>11,12,21,22
Shokri A. <i>et al.</i> [57]	2014	Iran	Cross-sectional	Radiographs (panoramic)	7-35	1649	5.7	Not specified		12,22>35,45>15,25
Bozga A <i>et al.</i> [58]	2014	Romania	Cross-sectional	Pre-orthodontic records	6-41	518	6.7	3.3	3.4	35,45 >12,22 >15,25 > 31,41 > 37,47 >32,42
Gonçalves-Filho AS <i>et al.</i> [59]	2014	Brazil	Cross-sectional	Dental records	2-30	487	6.16	2.26	3.9	Not specified
Herrera Atoche JR <i>et al.</i> [60]	2014	Mexico	Retrospective	Pre-orthodontic records	9-20	690	5.82	2.09	3.73	35,45
Karadas M <i>et al.</i> [61]	2014	Turkey	Retrospective	Radiographs (panoramic)	8- 16	2722	3.89	1.51	2.38	12,22>35,45>31,41,15,25

(Table 3) contd....

Reference Study	Publication Year	Country	Study Type	Evaluation Method	Age Range (Years)	Total Sample Size	Hypodontia %	Distribution between the Genders %		Most Frequently Reported Hypodontia
								Male	Female	
Mani SA <i>et al.</i> [62]	2014	Malaysia	Retrospective	Radiographs (panoramic)	12-16	834	7.3	3.4	3.9	12,22 >15,25,35,45 >13,23
Al-Amiri A <i>et al.</i> [63]	2013	America	Cross-sectional (retrospective)	Pre-orthodontic records	Not specified	496	9.5	Not specified		35,45>12,22>15,25
Cunha MGM <i>et al.</i> [64]	2013	Brazil	Cross-sectional (retrospective)	Radiographs (panoramic)	4-12	523	8.9	2.1	6.8	Anodontia
Diab H. [65]	2013	Saudi Arabia	Cross-sectional (retrospective)	Radiographs (panoramic)	25.3-26.8	350	15.4	7.14	8.28	12,22,32,42>15,25,35,45>
Trakinienė G. <i>et al.</i> [66]	2013	Lithuania	Retrospective	Pre-orthodontic records	10-39	824	17.11	4.85	12.26	35,45
Campoy MD <i>et al.</i> [67]	2013	Portugal	Retrospective	Dental records	7-21	2888	6.1	Not specified		
Rathi MK <i>et al.</i> [68]	2013	Pakistan	Retrospective	Radiographs (panoramic)	Not specified	570	6.8	3.3	3.5	35,45
Afify AR. <i>et al.</i> [69]	2012	Saudi Arabia	Cross-sectional (retrospective)	Dental records	12-30	878	4.66	2.39	2.27	34,35,44,45>13,23
Sogra Y <i>et al.</i> [70]	2012	Iran	Retrospective	Radiographs (panoramic)	Not specified	1590	10	1.26	8.74	12,22>35,45
Asec Coelho <i>et al.</i> [71]	2012	Portugal	Retrospective	Radiographs (panoramic)	6-15	1438	8	3.9	4.1	35,45> 12, 22 > 15,25
Medina AC [72]	2012	Venezuela	Cross-sectional	Dental records	5-11	607	4.1	1.5	2.6	12,22 > 35,45 > 15,25 >32,42 > 37,47
Fnaish M M [73]	2011	Jordan	Prospective	Clinical and radiographic examinations	5-12	3600	8.8	3.4	5.4	35,45>15,25>12,22
Gupta SK <i>et al.</i> [74]	2011	India	Cross-sectional	Clinical examination, dental models, and radiographs	Not specified	1123	4.54	2.31	2.23	12,22>31,41>11,21,14,15,25,24
Kazanci F <i>et al.</i> [75]	2011	Turkey	Cross-sectional	Clinical examination, dental models, and radiographs	9-25	3165	4.5	1.3	3.2	12,22>35,45>31,41 >15,25
Behr M. <i>et al.</i> [76]	2011	Germany	Retrospective	Dental records	5-44	1353	12.6	6.3	6.3	35>45>22>12>15>25
Kim YH [77]	2011	Korea	Retrospective	Clinical examination, dental models, and radiographs	9-30	3055	11.3	3.7	7.6	35,45>32,42>15,25
Aktan AM <i>et al.</i> [78]	2010	Turkey	Retrospective	Radiographs (panoramic)	5 - 37	100,577	1.47	0.62	0.85	35,45>12,22
Vahid-Dastjerdi E <i>et al.</i> [79]	2010	Iran	Retrospective	Dental models, and radiographs (panoramic, intraoral periapical)	9-27	1751	9.1	4.2	4.9	12,22>15,25>32,42> 35,45
Celikoglu M <i>et al.</i> [80]	2010	Turkey	Retrospective	Radiographs (panoramic)	10 - 25	3341	4.6	1.5	3.1	12,22>35,45>31,41
Hashem AA <i>et al.</i> [81]	2010	Ireland	Retrospective	Clinical examination and radiographs (panoramic)	7-50	168	11.3	Not specified		35,45>15,25>12,22
Gomes RR [82]	2010	Brazil	Retrospective	Pre-orthodontic records	10-15.7	1049	6.3	2.5	3.8	12,22
CJ Chung <i>et al.</i> [83]	2008	Korea	Retrospective	Dental models, and radiographs (panoramic)	Not Specified	1622	11.2	5.5	5.7	32,42>35,45
Goya, H. A. <i>et al.</i> [84]	2008	Japan	Retrospective	Radiographs (panoramic)	3-17	2072	8.4	4.1	4.3	35,45 > 12, 22> 15,25
Endo <i>et al.</i> [85]	2006	Japan	Retrospective	Radiographs (panoramic)	5 -15	3358	8.5	3.2	5.3	35,45> 12, 22, 32, 42 > 15, 25
Gábris K <i>et al.</i> [86]	2006	Hungary	Retrospective	Radiographs (panoramic)	6-18	2219	14.69	Not specified		12,22>35,45>15,25 >31,41

(Table 3) contd....

Reference Study	Publication Year	Country	Study Type	Evaluation Method	Age Range (Years)	Total Sample Size	Hypodontia %	Distribution between the Genders %		Most Frequently Reported Hypodontia
								Male	Female	
Silva Meza R. [87]	2003	Mexico	Retrospective	Radiographs (panoramic)	9-20	668	2.7	Not specified		12, 22>35,45
Ng'ang'a RN and Ng'ang'a PM [88]	2001	Kenya	Retrospective	Radiographs (panoramic)	8-15	615	6.3	3.9	2.4	35,45>15,25>12,22
Backman B. and Wahlin Y.B. [89]	2001	Switzerland	Cross-sectional	Clinical examination and radiographs (bitewing, intraoral periapical, extraoral)	7	739	7.4	3.2	4.2	35,45 > 15,25

The prevalence of hypodontia observed in this study was 7.8%, a finding consistent with investigations from France (7.9%), Yemen (7.48%), Brazil (7.44%), Croatia (7.8%), Qatar (7.8%), Malaysia (7.3%), and Switzerland (7.4%) [14, 31, 32, 45, 51, 62, 89]. This notable uniformity across various geographic regions highlights the probable influence of shared genetic predispositions and environmental factors, such as familial history, dietary influences, and exposure to environmental teratogens, contributing to the manifestation of this condition. Nevertheless, the prevalence rate observed in the present study surpasses the estimates reported in the meta-analysis by Khalaf *et al.* [12], which determined the global prevalence of this anomaly to be 6.4% and the Asian prevalence to be 6.3%. The literature reviewed in the present study further underscores notable variability in the prevalence of this abnormality, with reported frequencies ranging from 0.28% in the Indian population to 17.11% in the Albanian population [19, 35]. This variation in prevalence rates can be attributed to several factors, including patients' ethnicity, sampling techniques, diagnostic criteria, chronological age, and sample size, as highlighted by Shimizu T and Maeda T [11]. The disparity in methodologies and sample sizes influencing the prevalence rate of hypodontia is exemplified by five distinct Iranian studies, where prevalence rates span from 1.1% to 10% [28, 34, 49, 57, 70, 79].

The present study identified a notable variation in the distribution of hypodontia between males and females across different tooth types. This finding challenges prior research that suggested an equivalent predisposition to hypodontia among genders. Studies by Kumar D *et al.* [23], Hugo Norberto Aragón *et al.* [27], and Behr *et al.* [76] reported no significant gender differences in hypodontia prevalence within populations from India, Argentina, and Germany, respectively. Similarly, investigations conducted in the UAE [39], Romania [58], and Pakistan [68] found no statistical association between gender and hypodontia occurrence. However, a meta-analysis by Polder *et al.* [13] indicated a higher prevalence of hypodontia among females, who were approximately 1.4 times more likely to exhibit the condition than males. Additional studies from France [14], Turkey [30], Brazil [32], Sudan [47], Qatar [51], Thailand [56], Lithuania [66], Iran [70], and Korea [77] have also highlighted gender-based variability in hypodontia prevalence.

The findings of the present study revealed a markedly higher prevalence of hypodontia in the maxillary arch (75.57%) compared to the mandibular arch (24.43%). This observation corroborates the conclusions drawn by Khalaf *et al.* [12] in their comprehensive meta-analysis, which similarly reported a greater propensity for hypodontia in the maxilla relative to the mandible. In contrast, investigations conducted within French [14], Thai [17], and Latvian [16] populations have demonstrated a higher prevalence of hypodontia in the mandibular arch.

Our investigation identified the maxillary lateral incisors as the most frequently affected teeth in cases of hypodontia. This finding aligns with previous studies conducted on populations from Egypt [24], Iran [28, 49, 57, 70, 79], Yemen [31], Qatar [51], Saudi Arabia [55, 65], Turkey [75, 61, 80], and Malaysia [62]. However, meta-analyses by Khalaf *et al.* [12] and Polder *et al.* [13] reported the mandibular second premolar as the most commonly affected tooth globally. Furthermore, in the present study, hypodontia involving mandibular incisors was notably rare or absent, contrasting with their higher prevalence in Chinese and Japanese populations as reported in prior literature [90, 91].

In the present study, hypodontia predominantly manifested in its mild form, with either one (60%) or two (30%) missing teeth. This finding aligns with meta-analyses by Khalaf *et al.* [12] and Polder *et al.* [13], which estimated the prevalence of mild forms of hypodontia to be the highest, accounting for 81.6% and 83% of total cases, respectively. Moreover, this trend is consistent with observations reported in studies by Goya *et al.* [84] and Endo *et al.* [85], further corroborating the predominance of mild form of hypodontia across diverse populations.

Consistent with the meta-analyses conducted by Polder *et al.* [13], the present study demonstrated a higher prevalence of unilateral hypodontia compared to its bilateral counterpart. However, this finding contrasts with studies conducted on Italian [44], Japanese [85, 85], and Mexican [87] populations, where bilateral hypodontia was either more prevalent or exhibited an equal prevalence to unilateral hypodontia. In the present study, bilateral hypodontia was predominantly observed in the maxillary lateral incisors, maxillary first premolars, and mandibular second premolars. These findings align with the observations of previous studies conducted by Gracco *et al.* [38], Goya *et al.* [84], and Kirzioglu [86], which reported bilateral hypo-

dontia involving the same teeth, albeit in a differing order of prevalence.

Given the functional and aesthetic implications of hypodontia, particularly in cases of multiple missing teeth, timely and strategic management is critical. A multidisciplinary approach involving orthodontists, oral surgeons, and prosthodontists is essential to ensure optimal outcomes. Preservation of alveolar bone is a key factor in treatment planning, often achieved through orthodontic strategies, such as delayed extraction of retained primary teeth and implant site switching, which help maintain bone volume and prepare sites for future implant placement [92]. In situations where bone loss has occurred, surgical interventions like bone grafting or distraction osteogenesis may be necessary [93]. Preventive orthodontic techniques, including space opening and the use of rigid bonded retainers, can further support alveolar ridge preservation and implant site stability. Individualized treatment planning and coordination among specialties are paramount, particularly as technological advancements continue to expand the possibilities in hypodontia management [94, 95].

There are some limitations in the research. While this study provides valuable epidemiological data on hypodontia in the Nepalese population, several limitations must be acknowledged. First, as a single-center study utilizing convenience sampling, the findings may not fully represent the entire Nepalese population. Second, the retrospective design relied solely on panoramic radiographs, without clinical examinations or genetic analysis, limiting insights into etiological factors. Third, the exclusion of individuals with syndromic conditions and prior orthodontic treatment may have introduced selection bias. Fourth, while the study assessed hypodontia prevalence and patterns, it did not evaluate its functional, esthetic, or psychosocial impact. Lastly, although associated dental anomalies were noted, their statistical significance and clinical correlations require further investigation.

CONCLUSION

This study presents a comprehensive assessment of the prevalence and pattern of non-syndromic hypodontia among the Nepalese population, adding a critical data point to the global body of literature on dental anomalies. With a prevalence of 7.8%, hypodontia appears to be relatively common within this subgroup, with distinct demographic and anatomical patterns. The condition was significantly more prevalent in males, particularly affecting the maxillary arch and most frequently involving the maxillary lateral incisors. Unilateral hypodontia was found to be more common than bilateral cases, and the majority of cases fell within the mild category, involving one or two missing teeth. These findings mirror many international studies yet also highlight regional variation, reinforcing the influence of ethnic, genetic, and environmental factors, as well as differences in sampling methodology and diagnostic criteria.

Despite key methodological strengths, including robust sample size and standardized radiographic protocols, the retrospective, single-center design and absence of clinical and genetic data limit both generalizability and etiological insight. These constraints underscore the need for multi-center, prospective investigations integrating clinical, radio-

graphic, and molecular diagnostics. Such studies are critical to refine early detection, optimize individualized treatment planning, and inform public health interventions aligned with population-specific risk profiles.

AUTHORS' CONTRIBUTIONS

The authors confirm their contribution to the paper as follows: S.D.: Study conception, design, and data collection; S.D., D.R., M.N.M.S.: Draft manuscript.

LIST OF ABBREVIATIONS

NSH	= Non-syndromic Hypodontia
TA	= Tooth Agenesis
DNA	= Deoxyribonucleic Acid
PAX9	= Paired Box Gene 9
MSX1	= Muscle Segment Homeobox 1
AXIN2	= Axis Inhibitor 2
EDA	= Ectodysplasin A
IRC	= Institutional Review Committee
SPSS	= Statistical Package for the Social Sciences
FDI	= Fédération Dentaire Internationale

ETHICS APPROVAL AND CONSENT TO PARTICIPATE

Ethical approval for this study was obtained from the Institutional Review Committee of Kathmandu University School of Medical Sciences (IRC-KUSMS), Nepal. Reference number: IRC-KUSMS-61/19.

HUMAN AND ANIMAL RIGHTS

All procedures performed in studies involving human participants were in accordance with the ethical standards of institutional and/or research committee and with the 1975 Declaration of Helsinki, as revised in 2013.

CONSENT FOR PUBLICATION

Informed consent was waived for this retrospective study due to the exclusive use of de-identified patient data, which posed no potential harm or impact on patient care.

STANDARDS OF REPORTING

STROBE guidelines were followed.

AVAILABILITY OF DATA AND MATERIALS

The data supporting the findings of this article is available in the "Zenodo Repository" at "<https://doi.org/10.5281/zenodo.15364185>", reference number "15364185".

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CONFLICT OF INTEREST

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

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