



Epidemiological profile of children and adolescents with oral cleft treated at a referral center in Curitiba, PR, Brazil

Perfil epidemiológico da criança e do adolescente com fissura oral atendidos em um centro de referência em Curitiba, PR, Brasil

ALINNE VILLELA VENDRAMIN¹
CAMILLA LUIZA VON STEINKIRCH¹
GABRIELA TUBINO CZARNOBAY¹
MARIA GABRIELA REGO MONTANHA
REBELLO¹
WILLIAM SKRONSKI TON¹
MARCO AURÉLIO GAMBORGI²
RENATO NISIHARA^{1*}

Institution: Universidade Positivo,
Curitiba, PR, Brazil.

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■ ABSTRACT

Introduction: Oral cleft is the second major cause of congenital anomalies and represents a major craniofacial alteration in live births. The objective of this study was to analyze the epidemiological data collected from the Center for Comprehensive Care to Individuals with Cleft Lip and Palate in the period from January 2011 to December 2014. **Methods:** This retrospective study evaluated 1,262 medical records of patients with an oral cleft. After applying the inclusion and exclusion criteria, 52.7% of the medical records were included in the study. **Results:** Among the 666 medical records, 57.4% were of male patients and 42.6% were of female patients. Of these, 34.8% of the patients had a trans-foramen cleft, 27.2% had a pre-foramen cleft, 25.8% had a post-foramen cleft, and 12.2% had another type of cleft. Patients from Curitiba and the metropolitan region constituted 36.6% of the cases, and patients from rural areas of Paraná represented 61% of the visits to the care center. The median age at the first visit of the patients from Curitiba/metropolitan region and rural areas of Paraná was 1 and 2 months, respectively. The first surgery was performed at the care center at the age of 6 months in patients from Curitiba and metropolitan region and 7 months in patients from rural areas of Paraná. **Conclusion:** There was a predominance of boys and a higher prevalence of incomplete post-foramen clefts in the total population. Despite the long distance to the care center, children from rural areas of Paraná underwent the correction surgery and were treated at the referral center with an age difference of only 1 month compared with patients who lived in Curitiba, where the care center is located.

Keywords: Epidemiology; Cleft lip; Reconstructive surgery.

¹ Universidade Positivo, Curitiba, PR, Brazil.

² Hospital do Trabalhador, Curitiba, PR, Brazil.

■ RESUMO

Introdução: A fissura oral é a segunda maior causa de anomalias congênitas e representa a principal alteração craniofacial em nascidos vivos. O objetivo do presente estudo foi determinar os dados epidemiológicos do Centro de Atenção Integral ao Fissurado Labiopalatal, no período entre janeiro de 2011 e dezembro de 2014. **Métodos:** Estudo retrospectivo utilizando prontuários clínicos. Foram avaliados 1262 prontuários de pacientes portadores de fissura oral. Após aplicação dos critérios de inclusão e exclusão, 52,7% prontuários foram incluídos no estudo. **Resultados:** Entre os 666 prontuários, 57,4% foram do gênero masculino e 42,6% do feminino. Verificou-se que 34,8% dos pacientes apresentaram fissuras transforame, 27,2% fissuras pré-forame, 25,8% fissuras pós-forame e 12,2% outros tipos de fissuras. Pacientes oriundos de Curitiba e Região Metropolitana correspondem a 36,6%, aqueles do Interior do Paraná abrangem 61% dos atendimentos no Centro de Atenção. As medianas de idade na primeira consulta, entre os pacientes de Curitiba e Região Metropolitana e do Interior do Paraná, são de 1 mês e 2 meses, respectivamente. E a primeira cirurgia, realizada no Centro de Atenção, foi em torno de 6 meses, nos pacientes de Curitiba e Região Metropolitana, e de 7 meses naqueles oriundos do Interior do Paraná. **Conclusão:** Verificou-se predomínio de fissuras em meninos e maior frequência da fissura pós-forame incompleta. Observou-se que, apesar da distância, as crianças oriundas do Interior do Paraná realizaram a cirurgia de correção e chegaram ao centro de referência com apenas um mês de diferença em relação aquelas da cidade sede do Centro de Atenção Integral ao Fissurado Labiopalatal.

Descritores: Epidemiologia; Fenda labial; Procedimentos cirúrgicos reconstrutivos.

INTRODUCTION

An oral cleft (OC) develops unevenly on the face during the embryonic and fetal periods¹ and is the second major cause of congenital anomalies in live births^{2,3}. OC is clinically characterized by nonclosure of the lip, palate, or both and affects approximately 1 in 700 live births⁴. In Brazil, the exact number of subjects with OCs is unknown, but it is estimated that 180,000 individuals are affected by this anomaly, and mortality in the first year of life is approximately 35%^{5,6}. Approximately 10% of the cases have other associated anomalies, characterizing different syndromes.

The Center for Comprehensive Care to Individuals with Cleft Lip and Palate (Centro de Atenção Integral ao Fissurado Labiopalatal-CAIF) is a care center of the State Health Secretariat of Paraná (Secretaria de Estado da Saúde do Paraná-SESA) and a member of the Reference Network for the Treatment of Craniofacial Deformities, an institution created in 1998 by the federal government for accreditation of care services specializing in the treatment of craniofacial deformities⁷. All treatment modalities are provided by

the Unified Health System (SUS), social contributions, and contributions from nongovernmental organizations (NGOs), including Operation Smile, which is an international NGO that recruits volunteer physicians to operate on individuals with cleft lip and palate at no charge⁸. Therefore, comprehensive and multidisciplinary treatment is available to patients with OCs⁶.

OC has a negative impact on speech, hearing, physical appearance, and cognition and affects the health and social integration of individuals with OCs. OC causes high morbidity, emotional distress, and can lead to social exclusion⁹. This anomaly is caused by genetic and environmental factors, which can act in isolation or together. The treatment of OCs is multiprofessional and should be integrated, comprehensive, and continuous, starting at the age of 3 months^{9,10}.

Knowledge of the epidemiological profile of patients with OC treated at a specialized center is essential for the community, allowing evaluation of effectiveness of care services, enabling better structuring of these services, and subsidizing new research.

OBJECTIVE

The objective of this study was to analyze the epidemiological data obtained from the CAIF - a referral center in Curitiba, Paraná, Brazil - which treats patients from the state of Paraná and 17 other states, and evaluate the epidemiological profile of patients with cleft lip and/or cleft palate admitted to this center.

METHODS

This descriptive, cross-sectional, and retrospective study was approved by the Research Ethics Committee of our institution under Certificate for Ethics Assessment (Certificado de Apresentação para Apreciação Ética-CAAE) No. 44484015.0.0000.0093.

In total, 1,262 medical records of patients with OC were analyzed, and the records were subdivided into four age groups - infants (aged 0 to 2 years), pre-school children (aged 2 to 4 years), schoolchildren (aged 4 to 10 years), and adolescents (aged 11 to 19 years)¹¹, treated at the CAIF, located at the Worker's Hospital in Curitiba, Paraná.

The following variables were searched in the medical records: age, sex, place of origin, type of cleft (according to the Victor Spina classification¹²), age at the first visit, age at the first surgical intervention, and history of previous surgery. The inclusion criteria were cleft lip, cleft palate, or cleft lip and palate, with a follow-up at the CAIF from January 2011 to December 2014. The exclusion criteria were age over 19 years, incomplete and/or illegible medical records, and any other craniofacial anomaly.

The classification of clefts most commonly used in CAIF is that of Victor Spina. This classification takes into account the incisive foramen, the border between the primary and secondary palates, and allows the separation of clefts into three types: pre-foramen (cleft lips, which may be categorized into unilateral, bilateral, or median), post-foramen (cleft palates, which may be complete, involving the entire hard palate, or incomplete, involving only the uvula or a portion of the palate), and trans-foramen (unilateral or bilateral, involving the lip, dental arch, and whole palate)^{12,13}.

After applying these criteria, 408 medical records were excluded because they involved patients with craniofacial anomalies, 20 medical records because they were incomplete and/or illegible, and 168 records because the patients were older than 19 years. In total, 666 medical records were included in the present study, corresponding to 52.7% of all records analyzed.

Statistical analysis

The data were tabulated and expressed as medians, means, and standard deviations or as

frequencies and percentages. Statistical analysis was conducted using the statistical package Prism version 5.0 (GraphPad, San Diego, CA, USA) using the Kolmogorov-Smirnov test to assess data normality. Categorical variables were expressed as percentages and compared using the χ^2 test or Fisher's exact test, as appropriate. A *p*-value of less than 5% was considered statistically significant.

RESULTS

The demographic characteristics and classification of cleft lips and palates in children treated at the CAIF during the study period are shown in Table 1. It was not possible to include a group from other regions in the statistical analysis because of the low number of patients (*n* = 16).

In the present study, the number of male patients with OCs was significantly higher than that of female patients (*p* < 0.0001). In both sexes, incomplete post-foramen cleft was the most common anomaly (25.8%), including 74 cases in male patients and 72 cases in female patients, followed by left unilateral trans-foramen cleft, which occurred in 59 boys and 39 girls. Among the participants, 137 patients (20.6%) had a history of OC in the family, 471 (70.7%) had a history of no OC in the family, and 58 (8.7%) had no history.

The medical records of 106 patients were not considered for the statistical analysis in the interval between the first consultation and the first surgery performed at the CAIF because these records were incomplete or the patients had not yet undergone corrective surgery by the date of the survey. Among these, 49 patients were from Curitiba or the metropolitan region, 50 patients were from rural areas of Paraná, and seven patients were from other geographical regions.

The median age at the first visit to CAIF was higher in patients from rural areas (*p* = 0.0001) than that of patients living in Curitiba or the metropolitan region (Table 2 and Figure 1).

There was no significant difference in the interval between the first consultation and the surgical procedure performed at the CAIF (*p* = 0.91) in the patients from the capital and those from rural areas of Paraná (Table 2 and Figure 2).

DISCUSSION

An OC limits many aspects of the patient's life, including speech, hearing, and cognition, and hinders the interaction of the patient with social groups. However, this effect can often be reversed when interventions are performed early¹⁴. OC needs to be diagnosed early to ensure that the treatment follows established guidelines and achieves the best possible

Table 1. Demographic profile and classification of cleft lip and palate in children treated at CAIF from January 2011 to December 2014.

Variables	n	%	
Sex⁺			
Male	382	57.4	
Female	284	42.6	
Place of origin			
Curitiba and metropolitan area	244	36.6	
Rural areas of Paraná	406	61	
Other regions	16	2.4	
Classification of cleft			
Pre-foramen* (27.2%)	Complete unilateral	45	6.7
	Complete bilateral	6	0.9
	Incomplete unilateral	114	17.1
	Incomplete bilateral	16	2.4
Trans-foramen* (34.8%)	Left unilateral	98	14.7
	Right unilateral	58	8.7
	Bilateral	76	11.4
Post-foramen* (25.8%)	Complete	26	14.7
	Incomplete	146	22
Others (12.2%)	Other combinations	7	1
	Submucosal	49	7.4
	Mixed	19	2.9
	Rare*	6	0.9

* according to the classification of Victor Spina (<http://ctmc.lusiada.br/malformacoes-craniofaciais/>). ⁺ $p < 0.0001$ using the χ^2 test. CAIF: Centro de Atenção Integral ao Fissurado Labiopalatal (Center for Comprehensive Care to Individuals with Cleft Lip and Palate).

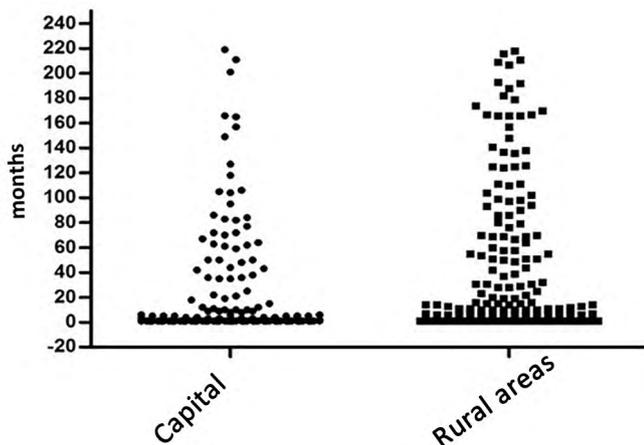
Table 2. Correlation between the place of origin, age (in months) at the first consultation at CAIF, age at the first surgery performed at CAIF, interval between the first consultation and first surgery performed at CAIF, and difference between patients previously operated or not.

Origin	Curitiba and metropolitan area, n = 244 (36.6%)	Rural areas of Paraná, n = 406 (61%)	Other regions, n = 16 (2.4%)
Age at first consultation at CAIF (months), median (range)	1 (1-219)	2 (1-218)	6 (1-98)
Age at the first surgical intervention at CAIF (months), median (range)	6 (1-222)	7 (2-222)	12 (4-81)
Interval between the first consultation and the first surgery performed at CAIF (months), median (range)	4 (1-41)	4 (1-24)	8 (3-15)
Previously operated (n = 54, 8.1%)	18	34	2
Not previously operated (n = 612, 91.9%)	226	372	14

CAIF: Centro de Atenção Integral ao Fissurado Labiopalatal (Center for Comprehensive Care to Individuals with Cleft Lip and Palate).

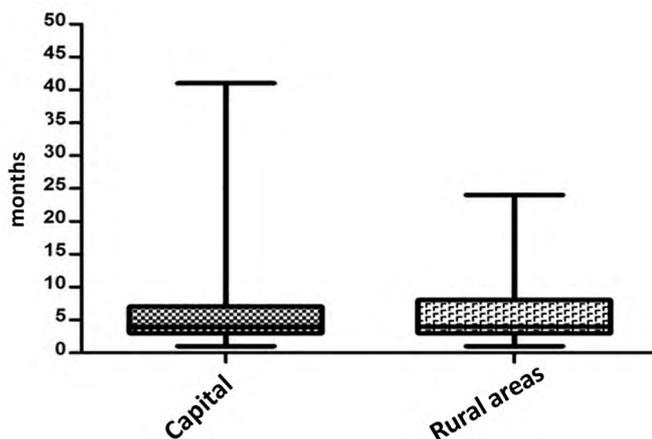
quality of life for the affected individuals¹⁵. Therefore, this study presented epidemiological data to evaluate the contribution of CAIF in the state of Paraná, to collect and disseminate data from this population, and indicate the importance of these data for the identification and effective treatment of children with cleft lip and palate.

The interval between the first consultation and the first surgery needs to be determined to ensure adequate planning of the scheduled surgical procedures. In fact, the earlier the patient is admitted to the CAIF, the higher the possibility of planning the surgery considering the recommended ages. This strategy aims at reducing the likelihood of hearing impairment and limitations to the child's maxillofacial



$p = 0.0001$, Mann-Whitney test.

Figure 1. Age (in months) of children at the first consultation at the Center for Comprehensive Care to Individuals with Cleft Lip and Palate (Centro de Atenção Integral ao Fissurado Labiopalatal-CAIF). The median age at the first visit to the CAIF was higher in patients from rural areas than that of patients living in Curitiba or the metropolitan region.



$p = 0.91$, Mann-Whitney test.

Figure 2. Period (in months) between the first visit and surgery considering the place of residence of patients treated at the Center for Comprehensive Care to Individuals with Cleft Lip and Palate (Centro de Atenção Integral ao Fissurado Labiopalatal-CAIF). There was no significant difference between the groups.

growth. Therefore, this interval should not be considered an absolute value without taking the age recommendations into account.

Regarding the age of the patient at admission to the care service, the most prevalent age group was infants, with a median age of 1 month among the patients from Curitiba and 2 months among the patients from rural areas of the state. A study conducted from 2006 to 2008 in Belo Horizonte¹⁶ and data obtained from a care service for patients with OCs located in the Northeast region⁹ indicated similar results regarding the prevalence of infants with OCs.

In the present study, a significant difference in the median age of children from the capital and children from rural areas at the first consultation at

the CAIF was observed, which can be attributed to the geographical distance to the referral center and the consequent displacement of patients coming from other regions to the center, which is located in the capital. However, even in cases in which treatment was provided at a median age of 2 months, the patients from rural areas were treated within the appropriate timeframe for the surgical correction, and the CAIF was effective even for these patients because of the promptness in performing the surgical procedure.

The recommended sequence of surgeries involves closure of the soft palate at the age of 3 to 9 months, depending on the type and extent of the cleft, and secondary closure of the residual hard palate at the age of 15 to 18 months. However, there is no consensus on these age intervals^{10,17}. In the present study, the age at which the CAIF received the patients with OCs was adequate considering the main objectives, which were ensuring speech development and maintaining normal maxillofacial growth in addition to reducing the incidence of otitis media and subsequent long-term hearing loss¹⁸.

In this study, surgical correction was performed at a median of 6 and 7 months in the patients from the capital and rural areas, respectively. Patients from rural areas had more reasons for nonadherence or might need to wait longer for surgery. However, our results indicated that the longer distance from the CAIF did not represent a significant barrier for this group to attend the visits scheduled for surgical procedures (Figure 2), indicating adequate performance of this referral center in the treatment of patients with OCs.

The number of cases of OC in male patients was higher than that in female patients. This result is similar to that found in other studies conducted in Brazil in São José dos Campos, São Paulo¹⁹, Recife, Pernambuco⁹, and Minas Gerais²⁰⁻²² and in Africa²³. Moreover, pre-foramen clefts were more common in boys (11.1%), but this result disagrees with those in the literature^{2,24}. The incidence of trans-foramen clefts was similar to that of other studies, with a lower incidence in girls.

Many studies reported a higher incidence of left unilateral clefts²⁵⁻³⁴, which contradicts the results of a study conducted in Iran³⁵ between 1982 and 2011, wherein the most common anomaly was bilateral followed by right unilateral clefts. However, the results of the remaining studies agree with ours, with a higher prevalence of left unilateral clefts (30.7%) followed by right unilateral clefts (16.5%) and bilateral clefts (14.7%). These results indicate the influence of environmental and genetic factors on the incidence and type of OC in different populations.

Epidemiological studies on OC use two common classifications of clefts. One was used in the present

study, and the other classification simplifies the malformations into cleft lip, cleft palate, and cleft lip and palate. We chose the Victor Spina classification because it was more commonly used in the referral center used for data collection and because it was more comprehensive in the details of the clefts. In our study as well as in other Brazilian studies^{2,36}, there was a predominance of trans-foramen clefts and cleft lips and palates. In a study conducted in Africa²³ and another in Iran³⁵, the most common type of malformation was combined cleft lip and palate, occurring in 58% and 50% of patients, respectively.

It was not possible to evaluate the ethnicity of the individuals with OCs treated at the CAIF. This information was not included in several medical records, thus limiting the analysis. Croen et al.²⁴ evaluated a population in California in the United States and observed that Black people had a lower prevalence of OC than Caucasians and that Asians had the highest prevalence, which agrees with the results of a study conducted in Europe²⁵.

The CAIF is an institution founded in 1993 and maintained by the SESA of Paraná with the assistance of the Association for Rehabilitation and Social Promotion of Individuals with Cleft Lips and Palates (Associação de Reabilitação e Promoção Social ao Fissurado Labiopalatal-AFISSUR). With a large, competent, and committed multidisciplinary team, this care center treats children and adults from 18 Brazilian states and partners with national and international organizations and universities. CAIF is recognized worldwide as a referral center for the treatment and rehabilitation of individuals with OCs and craniofacial deformities in general⁷.

CONCLUSION

In the present study, there was a predominance of male subjects with OCs. According to the classification of clefts, the most common deformity was incomplete post-foramen cleft. There was a significant difference in the median age at the first consultation between children living in the capital and those living in rural areas of Paraná. However, there was no difference in age at the time of surgery considering the place of origin of the patients. In addition, the interval between the first consultation and surgery was adequate with respect to what is expected from a referral center.

COLLABORATIONS

AVV Analysis and/or interpretation of data; final approval of the manuscript; conception and design of the study; completion of surgeries and/or experiments; writing the manuscript or critical review of its contents.

CLVS Final approval of the manuscript; conception and design of the study; completion of surgeries and/or experiments.

GTC Final approval of the manuscript; conception and design of the study; completion of surgeries and/or experiments.

MGRMR Final approval of the manuscript; conception and design of the study; completion of surgeries and/or experiments.

WST Final approval of the manuscript; conception and design of the study; completion of surgeries and/or experiments.

MAG Analysis and/or interpretation of data; final approval of the manuscript; conception and design of the study; completion of surgeries and/or experiments; writing the manuscript or critical review of its contents.

RN Analysis and/or interpretation of data; final approval of the manuscript; conception and design of the study; completion of surgeries and/or experiments; writing the manuscript or critical review of its contents.

REFERENCES

- Lemos IC, Feniman MR. Sustained Auditory Attention Ability Test (SAAAT) in seven-year-old children with cleft lip and palate. *Braz J Otorhinolaryngol*. 2010;76(2):199-205.
- Souza J, Raskin S. Estudo clínico e epidemiológico de fissuras orofaciais. *J Pediatr (Rio J)*. 2013;89(2):137-44. DOI: <http://dx.doi.org/10.1016/j.jpmed.2013.03.010>
- Wantia N, Rettinger G. The current understanding of cleft lip malformations. *Facial Plast Surg*. 2002;18(3):147-53. DOI: <http://dx.doi.org/10.1055/s-2002-33061>
- Dixon MJ, Marazita ML, Beaty TH, Murray JC. Cleft lip and palate: understanding genetic and environmental influences. *Nat Rev Genet*. 2011;12(3):167-78. DOI: <http://dx.doi.org/10.1038/nrg2933>
- Loffredo LDCM, Souza JMP, Yunes J, Freitas JAS, Spiri WC. Fissuras lábio-palatais: estudo caso-controle. *Rev Saúde Pública*. 1994;28(3):213-7. DOI: <http://dx.doi.org/10.1590/S0034-89101994000300009>
- Thomé S. Estudo da prática do aleitamento materno em crianças portadoras de malformação congênita de lábio e/ou de palato [Dissertação de mestrado]. Ribeirão Preto: Universidade de São Paulo. Escola de Enfermagem de Ribeirão Preto; 1990.
- Secretaria da Saúde. Centro de atendimento integral ao fissurado lábio palatal, institucional. Secretaria da Saúde. Paraná, Brasil. [acesso 2015 Ago 20]. Disponível em: <http://www.caif.saude.pr.gov.br/>
- Operação sorriso. Sobre nós: operação sorriso Brasil. [acesso 2015 Set 15]. Disponível em: <http://www.operacaosorriso.org.br/>
- Coutinho ALF, Lima MC, Kitamura MAP, Ferreira Neto J, Pereira RM. Perfil epidemiológico dos portadores de fissuras orofaciais atendidos em um Centro de Referência do Nordeste do Brasil. *Rev Bras Saúde Matern Infant*. 2009;9(2):149-56. DOI: <http://dx.doi.org/10.1590/S1519-38292009000200004>
- Fernandes JW. *Cirurgia Plástica. Bases e Refinamentos*. 2a ed. Curitiba: Primax Gráfica; 2012. p. 225-8.

11. Calendário de puericultura. Sociedade Brasileira de Pediatria; 2014. [acesso 2015 Ago 20]. Disponível em: http://www.sbp.com.br/fileadmin/user_upload/pdfs/CalendarioPuericultura_Jan2014.pdf
12. Spina V, Psillakis JM, Lapa FS, Ferreira MC. Classification of cleft lip and cleft palate. Suggested changes. *Rev Hosp Clin Fac Med Sao Paulo*. 1972;27(1):5-6. PMID: 4671376
13. Araruna R da C, Vendruscolo DM. Nutrition of children with cleft lip and cleft palate, a bibliographic study. *Rev Lat Am Enfermagem*. 2000;8(2):99-105.
14. Amstalden-Mendes LG, Xavier AC, Antunes DK, Ferreira ACRG, Tonocchi R, Fett-Conte AC, et al. Estudo multicêntrico da época do diagnóstico de fendas orais. *J Pediatr (Rio J)*. 2011;87(3):225-30.
15. Brasil. Ministério da saúde. Assistência de média e alta complexidade no SUS / Conselho Nacional de Secretários de Saúde. Brasília: CONASS; 2007. [acesso 2015 Abr 17]. Disponível em: http://bvsmms.saude.gov.br/bvs/publicacoes/colec_progestores_livro9.pdf
16. Di Ninno CQMS, Fonseca LFN, Pimenta MVE, Vieira ZG, Fonseca JA, Miranda ICC, et al. Epidemiological survey of patients with cleft lip and/or palate at a specialized center in Belo Horizonte, Brazil. *Rev CEFA*. 2011;13(6):1002-8.
17. Rohrich RJ, Love EJ, Byrd HS, Johns DF. Optimal timing of cleft palate closure. *Plast Reconstr Surg*. 2000;106(2):413-21; quiz 422; discussion 423-5. DOI: <http://dx.doi.org/10.1097/00006534-200008000-00026>
18. Zambonato TC, Feniman MR, Blasca WQ, Lauris JR, Maximino LP. Profile of patients with cleft palate fitted with hearing AIDS. *Braz J Otorhinolaryngol*. 2009;75(6):888-92. DOI: <http://dx.doi.org/10.1590/S1808-86942009000600020>
19. Cerqueira MN, Teixeira SC, Naressi SCM, Ferreira APP. Occurrence of lip and palate clefts in the city of São José dos Campos-SP. *Rev Bras Epidemiol*. 2005;8(2):161-6. DOI: <http://dx.doi.org/10.1590/S1415-790X2005000200008>
20. Martelli Júnior H, Orsi Júnior J, Chaves MR, Barros LM, Bonan PRF, Freitas JAS. Estudo epidemiológico das fissuras labiais e palatais em Alfenas-Minas Gerais-de 1986 a 1998. *RPG Rev*. 2006;13(1):31-5.
21. Paranaíba LM, Almeida Hd, Barros LM, Martelli DR, Orsi Júnior JD, Martelli Júnior H. Current surgical techniques for cleft lip-palate in Minas Gerais, Brazil. *Braz J Otorhinolaryngol*. 2009;75(6):839-43. DOI: [http://dx.doi.org/10.1016/S1808-8694\(15\)30546-2](http://dx.doi.org/10.1016/S1808-8694(15)30546-2)
22. Martelli DR, Machado RA, Swerts MS, Rodrigues LA, Aquino SN, Martelli Júnior H. Non syndromic cleft lip and palate: relationship between sex and clinical extension. *Braz J Otorhinolaryngol*. 2012;78(5):116-20. PMID: 23108830
23. Conway JC, Taub PJ, Kling R, Oberoi K, Doucette J, Jabs EW. Ten-year experience of more than 35,000 orofacial clefts in Africa. *BMC Pediatr*. 2015;15:8. DOI: <http://dx.doi.org/10.1186/s12887-015-0328-5>
24. Croen LA, Shaw GM, Wasserman CR, Tolarová MM. Racial and ethnic variations in the prevalence of orofacial clefts in California, 1983-1992. *Am J Med Genet*. 1998;79(1):42-7. PMID: 9738868 DOI: [http://dx.doi.org/10.1002/\(SICI\)1096-8628\(19980827\)79:1<42::AID-AJMG11>3.0.CO;2-M](http://dx.doi.org/10.1002/(SICI)1096-8628(19980827)79:1<42::AID-AJMG11>3.0.CO;2-M)
25. Derijcke A, Eerens A, Carels C. The incidence of oral clefts: a review. *Br J Oral Maxillofac Surg*. 1996;34(6):488-94. PMID: 8971440 DOI: [http://dx.doi.org/10.1016/S0266-4356\(96\)90242-9](http://dx.doi.org/10.1016/S0266-4356(96)90242-9)
26. Stoll C, Alembik Y, Dott B, Roth MP. Associated malformations in cases with oral clefts. *Cleft Palate Craniofac J*. 2000;37(1):41-7. PMID: 10670888 DOI: [http://dx.doi.org/10.1597/1545-1569\(2000\)037<0041:AMICWO>2.3.CO;2](http://dx.doi.org/10.1597/1545-1569(2000)037<0041:AMICWO>2.3.CO;2)
27. Freitas JA, Dalben Gda S, Santamaria M Jr, Freitas PZ. Current data on the characterization of oral clefts in Brazil. *Braz Oral Res*. 2004;18(2):128-33. DOI: <http://dx.doi.org/10.1590/S1806-83242004000200007>
28. Shapira Y, Lubit E, Kuftinec MM, Borell G. The distribution of clefts of the primary and secondary palates by sex, type, and location. *Angle Orthod*. 1999;69(6):523-8. PMID: 10593442
29. Rajabian MH, Sherkat M. An epidemiologic study of oral clefts in Iran: analysis of 1,669 cases. *Cleft Palate Craniofac J*. 2000;37(2):191-6. PMID: 10749061 DOI: [http://dx.doi.org/10.1597/1545-1569\(2000\)037<0191:AESOC>2.3.CO;2](http://dx.doi.org/10.1597/1545-1569(2000)037<0191:AESOC>2.3.CO;2)
30. Murray JC, Daack-Hirsch S, Buetow KH, Munger R, Espina L, Paglinawan N, et al. Clinical and epidemiologic studies of cleft lip and palate in the Philippines. *Cleft Palate Craniofac J*. 1997;34(1):7-10. PMID: 9003905 DOI: [http://dx.doi.org/10.1597/1545-1569\(1997\)034<0007:CAESOC>2.3.CO;2](http://dx.doi.org/10.1597/1545-1569(1997)034<0007:CAESOC>2.3.CO;2)
31. Bellis TH, Wohlgenuth B. The incidence of cleft lip and palate deformities in the south-east of Scotland (1971-1990). *Br J Orthod*. 1999;26(2):121-5. PMID: 10420246 DOI: <http://dx.doi.org/10.1093/ortho/26.2.121>
32. Chuangsuwanich A, Aojanepong C, Muangsombut S, Tongpiew P. Epidemiology of cleft lip and palate in Thailand. *Ann Plast Surg*. 1998;41(1):7-10. PMID: 9678461 DOI: <http://dx.doi.org/10.1097/0000637-199807000-00002>
33. Collares MVM, Westphalen ACA, Costa TCD, Goldim JR. Fissuras lábio-palatinas: incidência e prevalência da patologia no Hospital de Clínicas de Porto Alegre: um estudo de 10 anos. *Rev AMRIGS*. 1995;39(3):183-8.
34. Cooper ME, Stone RA, Liu Y, Hu DN, Melnick M, Marazita ML. Descriptive epidemiology of nonsyndromic cleft lip with or without cleft palate in Shanghai, China, from 1980 to 1989. *Cleft Palate Craniofac J*. 2000;37(3):274-80. PMID: 10830807 DOI: [http://dx.doi.org/10.1597/1545-1569\(2000\)037<0274:DEONCL>2.3.CO;2](http://dx.doi.org/10.1597/1545-1569(2000)037<0274:DEONCL>2.3.CO;2)
35. Kianifar H, Hasanzadeh N, Jahanbin A, Ezzati A, Kianifar H. Cleft lip and Palate: A 30-year Epidemiologic Study in North-East of Iran. *Iran J Otorhinolaryngol*. 2015;27(78):35-41.
36. Gardenal M, Bastos PRHO, Pontes ERJC, Bogo D. Prevalência das fissuras orofaciais diagnosticadas em um serviço de referência em casos residentes no estado de Mato Grosso do Sul. *Arq Int Otorrinolaryngol*. 2011;15(2):133-41. DOI: <http://dx.doi.org/10.1590/S1809-48722011000200003>

Corresponding author:*Renato Nishihara**

Rua Prof. Pedro Viriato Parigot de Souza, 5300 - Cidade Industrial - Curitiba, PR, Brazil

Zip Code 81280-330

E-mail: renatonishihara@gmail.com