

DATA NOTE

Open Access



A cohort study among 402 patients with penile cancer in Maranhão, Northeast Brazil with the highest worldwide incidence

Ciro Bezerra Vieira¹, Antonio Teixeira-Júnior², Laisson Feitoza³, Jaqueline Pinho⁴, José Calixto⁵, Francisco Sérgio Moura Silva do Nascimento¹, Marcos Adriano Garcia Campos¹, Joyce Lages¹, Antonio Machado Alencar Junior¹, Fernando Soares⁶, Isabela Cunha⁷ and Gyl Eanes Barros Silva^{4,8*} 

Abstract

Objective: Maranhão State—Northeast Brazil—has the world's highest incidence of penile cancer. This study describes the epidemiological, histopathological and clinical profile of patients stricken across that Brazilian state. The study is aimed at providing new data on neoplasia.

Data description: 402 men stricken with penile cancer were studied from January 2004 to December 2018. A retrospective stage was developed with collection of physical and electronic records. A prospective stage was performed with collection of clinical and epidemiological information through a questionnaire. The surgical material was looked into by a uropathologist, and the lesions were evaluated for macroscopic characteristics and various microscopic parameters. Three articles using this data have already been published.

Keywords: Carcinoma, Penile neoplasms, Papillomavirus infections, Squamous cell carcinoma

Objective

The incidence of penile cancer is low in developed countries; however, this neoplasm is still a problem in developing countries, reaching alarming levels [1]. The main risk factors are strongly related to socioeconomic conditions, genital hygiene habits, characteristics related to the foreskin and human papillomavirus (HPV) infection [2, 3]. The state of Maranhão—Northeastern Brazil—has the highest reported incidence of cancer of the penis (6.1 cases/100,000 ASR) and this fact is due to the high prevalence of risk factors in this region [4]. Maranhão is a poor state with highest prevalence of HPV infection in the country and has high rates of phimosis among the

male population [5, 6]. Penile cancer is a poorly studied topic compared to other more prevalent neoplasms, and few studies have been published on the population of the state of Maranhão. Thus, it is expected that the description of the characteristics of this population will contribute to the prevention and management of penile neoplasia, particularly with regard to the understanding of related risk factors with new information. These data sets are useful for comparing this population with that in other regions of the world and the significant number of cases analyzed also gives the study an important statistical significance. Few advances have been made in the management of penile neoplasia in recent years, and this study also aims to provide theoretical ground for further research of the diagnosis and treatment of this condition. Part of the data obtained in this study was used in three previously published studies [4, 5, 7], addressing the epidemiological and histopathological profile of penile cancer in Maranhão, as well as immunohistochemical

*Correspondence: gyleanes@fmrp.usp.br

⁸ Department of Pathology, Ribeirão Preto Medical School, University of São Paulo (USP), Bandeirantes Avenue, Monte Alegre, Ribeirão Preto 14049-900, Brazil

Full list of author information is available at the end of the article



analyzes related to the presence of HPV and of the p16INK4a protein in this sample.

Data description

This is a retrospective and prospective cohort study carried out in the state of Maranhão—Northeastern Brazil—from January 2004 to December 2018. The total population studied comprised 402 men with penile cancer treated by spontaneous demand at the Urology ward of two public hospitals within the state's capital city: University Hospital of the Federal University of Maranhão and Aldenora Bello Cancer Hospital. There were no exclusion criteria. The study variables included epidemiological, clinical and surgical characteristics of the participants and histopathological aspects of the penis cancer samples (see Table 1). The criterion used to classify genital hygiene as poor/moderate was less than one genital wash per day or the presence of phimosis [8]. The tumor subtype was described according to the revised and updated classification of penile carcinomas of the World Health Organization (2016) [9]. The histological grade of the tumor was described according to the classification by Velasquez et al. [10] and the staging was performed according to the classification of the TNM, 8th ed., 2017. Data collection was performed in two stages. First, all data corresponding to the period from January 2004 to December 2014 was collected retrospectively through physical and electronic records and amounted to 286 patients (see Table 1). From January 2015 to December 2018, the epidemiological and clinical data of 116 patients were prospectively collected through a questionnaire prepared by the research team (see Table 1); the questionnaire was applied by a single researcher directly to patients during outpatient consultations or during the period of hospital stay for surgical procedures. The surgical materials extracted from each patient were sent to the Laboratory of Immunofluorescence and Electron Microscopy at the University Hospital of the Federal University of Maranhão (LIME–HUUFMA). The lesions were photographed and those that were not included in paraffin were either completely embedded in paraffin

or made at least 40 paraffin blocks for rather advanced lesions. The histopathological material was examined by a single experienced uropathologist (GEBS) for macroscopic characteristics and the various microscopic parameters. Some samples of paraffin tumor tissue and fresh tissue were selected for immunohistochemistry and analysis of the polymerase chain reaction (PCR), respectively. Histological identification and molecular analysis of HPV, as well as immunohistochemistry for expression of p16INK4a were performed according to the description contained in the study by Martins VA et al. [7]. The population data obtained in the first stage of the study were used to calculate the age-standardized incidence rate (ASR) of penile cancer in Maranhão [4] with the very same population. Categorical variables were presented by means of frequencies and percentages and numerical variables by means and standard deviations (mean \pm SD). The data were analyzed using the Stata program, version 12.0. The association between categorical variables was analyzed using the Chi square test or Fisher's exact test. The 95% confidence interval was calculated by logistic regression with a significance of $p < 0.05$.

Limitations

This study has some limitations: 1. The retrospective collection of part of the data came up against the difficulty in obtaining complete information about some variables which were sometimes not recorded or were recorded in an unreadable way within the physical records; however, the data obtained were sufficient to understand important aspects of penile cancer in Maranhão region. 2. Data collection through a questionnaire, in some cases, ran into the difficulty of understanding by some patients with low education; nevertheless, in five cases in which the collection was considered insufficient, patients were excluded from the study. 3. There was loss of monitoring or abandonment of treatment in many cases, probably due to the fact that they live in distant rural areas, with difficulty in accessing the hospitals participating in the study.

Table 1 Overview of data files/data sets

Label	Name of data file/data set	File types (file extension)	Digital object identifier (DOI)
Data file 1	Retrospective profile of penile cancer in Maranhão (2004–2014)	.xlsx	Figshare (https://doi.org/10.6084/m9.figshare.12470045.v4) [11]
Data file 2	Prospective profile of penile cancer in Maranhão (2016–2018)	.xlsx	Figshare (https://doi.org/10.6084/m9.figshare.12470030.v4) [12]
Data file 3	P16 ^{INK4a} expression in patients with penile cancer	.xlsx	Figshare (https://doi.org/10.6084/m9.figshare.12921941.v1) [13]

Abbreviations

HPV: Human papillomavirus; STIs: Sexually transmitted infections; LIEM: Laboratory of Immunofluorescence and Electron Microscopy.

Acknowledgements

Not applicable.

Authors' contributions

CBV and GEBS were responsible for acquisition of data and drafting of the manuscript. ATJ and JC participated in the care and management of the patient and data collection. GEBS and IC performed pathological analysis and interpretation. JP and MAGC contributed to statistical analysis. JL, IC, FS, LF, FSMSN and AMAJ contributed to critical revision of important intellectual content of the manuscript. All authors contributed to the writing process and read and approved the final manuscript. All authors read and approved the final manuscript.

Funding

This study and 3 other researches about penile cancer in Maranhão developed by our research group were funded by the Research Support Foundation of Maranhão (Fundação de Amparo a Pesquisa do Maranhão—FAPEMA).

Availability of data and materials

The data described in this Data Note can be freely and openly accessed on the figshare dataset repositories in the following data records: [<https://doi.org/10.6084/m9.figshare.12470045.v4>], [<https://doi.org/10.6084/m9.figshare.12470030.v4>], [<https://doi.org/10.6084/m9.figshare.12921941.v1>]. Please see Table 1 and references [11–13] for details and links to the data.

Ethics approval and consent to participate

This study was approved by the Research Ethics Committee of the Universidade Federal do Maranhão [process no. 43,774,215.7.0000.5086], and written informed consent was obtained from all patients.

Consent for publication

Not applicable.

Competing interests

The authors declare that they have no competing interests.

Author details

¹ University Hospital of Federal University of Maranhão (UFMA), Barão de Itaparí Street, Centro, São Luís, Brazil. ² Postgraduate Program in Adult Health (PPGSAD), Federal University of Maranhão (UFMA), São Luís, MA, Brazil. ³ Department of Radiology, University Clinic Hospital of Estadual University of Campinas, Campinas, Brazil. ⁴ Laboratory of Immunofluorescence and Electron Microscopy (LIME), Presidente Dutra University Hospital (HUUFMA), São Luís, MA, Brazil. ⁵ Department of Medicine II, Federal University of Maranhão (UFMA), São Luís, MA, Brazil. ⁶ Department of Pathology, School of Medicine, University of São Paulo, São Luís, Brazil. ⁷ Doctor at the Antônio Prudente Foundation, São Paulo, Brazil. ⁸ Department of Pathology, Ribeirão Preto Medical School, University of São Paulo (USP), Bandeirantes Avenue, Monte Alegre, Ribeirão Preto 14049-900, Brazil.

Received: 8 July 2020 Accepted: 10 September 2020

Published online: 18 September 2020

References

- Christodoulidou M, Sahdev V, Houssein S, Muneer A. Epidemiology of penile cancer. *Curr Probl Cancer*. 2015;39(3):126–36.
- Barrasso R, De Brux J, Croissant O, Orth G. High prevalence of papillomavirus-associated penile in-traepithelial neoplasia in sexual partners of women with cervical intraepithelial neoplasia. *N Engl J Med*. 1987;317:916–23.
- Maden C, Sherman KJ, Beckmann AM, Hislop TG, Teh CZ, Ashley RL, et al. History of circumcision, medical conditions, and sexual activity and risk of penile cancer. *J Natl Cancer Inst*. 1993;85:19–24.
- Coelho RWP, Pinho JD, Moreno JS, et al. Penile cancer in Maranhão, Northeast Brazil: the highest incidence globally? *BMC Urol*. 2018;18:50.
- Vieira CB, Feitoza L, Pinho J, et al. Profile of patients with penile cancer in the region with the highest worldwide incidence. *Sci Rep*. 2020;10(1):2965.
- Estudo Epidemiológico sobre a Prevalência Nacional de Infecção pelo HPV (POP-Brasil): Resultados preliminares (Associação Hospitalar Moinhos de Vento, Porto Alegre, 2017).
- Martins VA, et al. P16INK4a expression in patients with penile cancer. *PLoS ONE*. 2018;13:e0205350. <https://doi.org/10.1371/journal.pone.0205350>.
- Chaux A, Netto GJ, Rodríguez IM, et al. Epidemiologic profile, sexual history, pathologic features, and human papillomavirus status of 103 patients with penile carcinoma. *World J Urol*. 2013;31(4):861–7.
- Cubilla AL, Velazquez EF, Amin MB, et al. The World Health Organization 2016 classification of penile carcinomas: a review and update from the International Society of Urological Pathology expert -driven recommendations. *Histopathology*. 2018;72(6):893–904.
- Velazquez EF, Ayala G, Liu H, Chaux A, Zanotti M, Torres J, et al. Histologic grade and perineural invasion are more important than tumor thickness as predictor of nodal metastasis in penile squamous cell carcinoma invading 5 to 10 mm. *Am J Surg Pathol*. 2008;32:974–9.
- Vieira, C. et al. Prospective profile of 116 men with penile cancer in Maranhão (2016–2018). *figshare*. <https://doi.org/10.6084/m9.figshare.12470045.v3> (2020).
- Vieira, C. et al. Retrospective profile of 286 men with penile cancer in Maranhão (2004–2014). *figshare*. <https://doi.org/10.6084/m9.figshare.12470030.v3> (2020).
- Martins, de Andrade. et al. P16^{INK4a} expression in patients with penile cancer. *figshare*. <https://doi.org/10.6084/m9.figshare.12921941.v1> (2020).

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Ready to submit your research? Choose BMC and benefit from:

- fast, convenient online submission
- thorough peer review by experienced researchers in your field
- rapid publication on acceptance
- support for research data, including large and complex data types
- gold Open Access which fosters wider collaboration and increased citations
- maximum visibility for your research: over 100M website views per year

At BMC, research is always in progress.

Learn more biomedcentral.com/submissions

