

# Differences in the diagnosis of primary cutaneous melanoma in the public and private healthcare systems in Joinville, Santa Catarina State, Brazil\*

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Abstract: Background: Cutaneous melanoma accounts for up to 80% of deaths caused by skin cancer. Diagnostic suspicion and access to medical care and early intervention in suspected cases is vital to the patient's prognosis.

OBJECTIVES: To compare demographic and histopathological characteristics of primary cutaneous melanoma diagnosed in the public healthcare system (Sistema Único de Saúde SUS) and the private system in Joinville, Santa Catarina State, Brazil.

METHODS: This cross-sectional retrospective study analyzed primary cutaneous melanoma cases recorded from 2003 to 2014 in the resident population of Joinville. Ethical approval was obtained from the local Research Ethics Committee.

RESULTS: 893 cases of primary cutaneous melanoma were identified. Patients in the private system were mostly younger, while there were more elderly patients in the public healthcare system (p <0.001). There was no statistically significant association between type of care (public/private) and gender or presence of multiple primary cutaneous melanomas. Histological diagnosis of superficial spreading melanoma was more common in patients treated in private healthcare, while nodular melanoma was more frequent in patients in the public healthcare system (p <0.001). Mean Breslow depth in patients treated in private healthcare was 1.35mm, compared to 2.72mm in the public system (p < 0.001).

STUDY LIMITATIONS: This was a retrospective study using secondary databases.

CONCLUSIONS: thin cutaneous melanoma (in situ cutaneous melanoma and Breslow T1) showed the strongest association with the private healthcare system, while thick cutaneous melanoma was more frequent in the public system (Breslow category T3 and T4) (p < 0.001).

Keywords: Health systems; Melanoma; Public health

# INTRODUCTION

Cutaneous melanoma (CM) accounts for up to 80% of skin cancer deaths.<sup>1,2</sup> The risk of developing metastatic melanoma and dying from CM is directly related to the phase of the disease at the time of diagnosis.<sup>3,4</sup> Survival rates are based on the TNM classification (which assesses thickness of the primary tumor and presence of distant or lymph node metastatic disease) and vary significantly according to these aggravating factors.5 Thus, timely detection, assessment and management should be a high public health priority, and the dermatologist plays a crucial role in this process.<sup>6-8</sup>

In Brazil, epidemiological data on CM are limited at both the national and regional levels.9 According to the Brazilian National Cancer Institute (INCA), the projected crude rates in 2016 were 3.03 cases/100,000 men and 2.59/100,000 women, with higher estimated rates in the South of Brazil and higher mortality in men. 10-11

In 2014, the municipality (county) of Joinville had 554,601 inhabitants, predominantly white (86%), with an important young contingent (30% ≤ 20 years of age).12 Joinville has a high CM incidence by Brazilian standards, with an upward trend and higher

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incidence in women. From 2003 to 2006, Joinville's CM age-standardized incidence (2010 World population standard) was 11.8 per 100,000 persons, and from 2011 to 2014 it was 17.5, an increase of 48.3% (p<0.05).<sup>13</sup>

Given this background, the study aimed to compare the demographic and histopathological characteristics of primary CMs diagnosed in patients in the public healthcare system (SUS) with those of private patients (health plans, health insurance, out-of-pocket) in Joinville. The purpose was to identify differences between these patient groups and thus help orient preventive measures, especially early diagnosis.

## **METHODS**

This retrospective cross-sectional study analyzed cases of primary CM recorded from January 2003 to December 2014 in the Joinville population. The study period was selected due to the availability of systematic records on CM in the laboratories.

Pathology reports were collected on all cases of primary CM diagnosed in the only three anatomical pathology laboratories in the municipality, which are responsible for diagnosis in patients residing in Joinville and the region. In two laboratories, data were collected with the descriptors "melanoma" and "malignant lentigo" in the field for diagnosis and the descriptor "skin and/or cutaneous" in the field "affected organ" to identify and select the reports for all CMs in the electronic databases. The third laboratory provided its diagnostic record books for the years 2003 to 2007 for manual collection, while in subsequent years the data were collected as in the first two laboratories.

Patients' municipality of residence was not available in half of the pathology reports. We thus searched for this information in the user database of the Unified Health System (SUS), under the responsibility of the Municipal Health Department, the database of patients covered by a health plan, and the place of origin of the specimen (specialty outpatient clinics of the SUS and private physicians' offices). Patients' full name and date of birth were used to confirm the address and to identify the existence of multiple pathology reports for the same CM (which can happen when the diagnosis is the result of an incisional biopsy followed by tumor excision or expansion of the margins). For patients with CM reports resulting from incisional biopsy and subsequent excision, the study used the report with the greatest Breslow depth. The study had no access to patients' medical charts or to the patients themselves. The histological specimens were not reviewed, assuming as correct the diagnosis and descriptions found in the pathology reports.

Inclusion criteria were residence in Joinville and having a diagnosis of primary CM during the study period. Exclusion criteria were: pathology reports on specimens from other cities; cases of melanoma of other organs and tissues; reports on revision of slides from surgical pathology; reports suggestive of melanoma, but with an inconclusive diagnosis, without immunohistochemical confirmation; reports on residual tumor or expansion of surgical margins.

After applying the inclusion and exclusion criteria to the 1,548 existing reports, 655 reports (42.3%) were excluded, as follows: reports on patients residing in other cities (390; 25.2%), melanoma metastases (46; 3.0%), duplicate reports (expansion of margins, inci-

sional biopsy followed by complete excision) (144; 9.30%), revision of sides or complementary immunohistochemistry (40; 2.6%), and other reasons (e.g., inconclusive reports, other neoplasms, melanomas of other organs) (35; 2.3%).

The procedures are believed to have included all cases of CM in patients residing in Joinville, making this a population-based study.

Data were keyed into Excel 2011 and later exported to SPSS v.18.0 for the statistical analysis. Categorical variables were described as frequencies and percentages and quantitative variables as mean with standard deviation, median, and interquartile range. Chi-square test was used to compare the categorical variables, and adjusted residuals analysis was used to locate the differences indicated by this test. Residuals with absolute values greater than 1.96 were considered statistically significant. Multivariate logistic regression was used to adjust for the age difference between the two patient groups (public versus private). Mann-Whitney test was used to compare the median Breslow depth. Statistical significance was set at 0.05.

The Breslow index, which assesses tumor thickness in millimeters, was categorized as T1 ( $\leq$ 1.0mm), T2 (1.01 – 2.0mm), T3 (2.01 – 4.0mm), T4 (>4.0mm), and not recorded or compromised.<sup>5</sup>

Patients were categorized by gender (female/male) and grouped by age bracket as young (under 60 years) or old (60 years or older), according to the WHO definition.<sup>14</sup>

The study was approved by the Institutional Review Board of Universidade da Região de Joinville, under case review no. 830.587.

# RESULTS

During the study period, 893 primary CMs were identified, with a predominance in women (507 or 56.8%). Mean age of CM patients was 54.6 years (SD=16.5). There was no statistical difference between the two groups (public versus private) according to gender (p=0.264). There was a predominance of younger patients in the private system and older patients in the public system (p<0.001). Individuals with multiple primary CMs predominated in the public system (p<0.001) (Table 1).

There was a statistically significant difference in histological type according to type of care, with superficial spreading melanoma (SSM) found more often in private patients (p<0.001) and nodular melanoma (NM) in patients in the public system (p<0.001) (Table 2).

Median Breslow depth in private patients was 0.75mm (interquartile range = 0.8850), compared to public patients with 1.30mm (interquartile range = 2.9075), a statistically significant difference (p<0.001).

Table 3 shows the statistically significant association between type of healthcare service and Breslow depth (p<0.001). According to adjusted residuals analysis, the strongest associations were between private care and *in situ* CM and Breslow T1, and between the public system and Breslow depths T3 and T4.

In the logistic regression analysis, the variables that showed an association with the type of healthcare system coverage were patient's age and histological level. Patients over 60 years of age had 1.64 times higher odds of treatment in the public healthcare system when compared to patients under 60 years, independently of the patient's histological level.

Patients with T1 tumors did not differ significantly from *in situ* tumors in relation to type of healthcare system coverage. Patients with T2 tumors had 1.75 times higher odds of treatment in the public healthcare system, compared to *in situ* patients. In patients with thicker tumors (Breslow T3 and T4), the odds of coming from the public system were more than double (3.79 and 3.80 times, respectively) when compared to *in situ* patients.

Logistic regression showed that the association between histological level and type of healthcare system coverage was independent of the patient's age. In other words, the association remained even after adjusting for age. Table 4 shows the adjusted logistic regression.

## DISCUSSION

In our study, 59% of individuals with CM were treated in private care (health plans, health insurance, out-of-pocket). This percentage is higher than the combined coverage rate for private health plans and health insurance as a whole in Joinville, which is 41% of the population according to the National Supplementary Healthcare Agency (ANS). This overall coverage, some 15% high-

TABLE 1: Cutaneous melanoma cases according to type of healthcare system (private or public) by gender, age bracket, and presence of multiple primary CMs in Joinville, Santa Catarina State, Brazil, 2003-2014

Type of healthcare system		Private		Public		p-value*
	N	N	(%)	N	(%)	
Total	893	526	(58.9)	367	(41.1)	
Gender						0.264
Female	507	290	(57.2)	217	(42.8)	
Male	386	236	(61.1)	150	(38.9)	
Age bracket (years)						< 0.001
Young (0-59)	546	352	(64.5)	194	(35.5)	
Old (≥ 60)	347	174	(50.1)	173	(49.9)	
Multiple CMs						0.999
Yes	127	52	(40.9)	75	(59.0)	
No	766	474	(61.9)	292	(38.1)	

Note: \* = Chi-square test.

Table 2: Distribution of CMs by histological type in patients treated in private versus public healthcare system (SUS)						
		Private		Public		
Histological type	N	N	(%)	N	(%)	
LMM	91	57	(10.9)	34	(9.4)	
DM	5	2	(0.4)	3	(0.8)	
ALM	10	5	(1.0)	5	(1.4)	
NMM	9	6	(1.1)	3	(0.8)	
NM	168	63	(12.1)	105	(28.9)*	
Melanoma, NOS	86	46	(8.8)	40	(11.0)	
SSM	516	343	(65.7)*	173	(47.7)	
Total**	885	522	(100.0)	363	(100.0)	

Note: Chi-square test - p-value <0.001; \* = adjusted residuals analysis; \*\* = missing data. LMM = lentigo maligna melanoma; DM = desmoplastic melanoma; ALM = acral lentiginous melanoma; NMM = nevoid malignant melanoma; NM = nodular melanoma; SSM = superficial spreading melanoma; NOS = not otherwise specified.

Table 3: Distribution of CMs in patients treated in private versus public healthcare system (SUS), according to histological level						
	_	Private		Public		
Histological type	n	n	(%)	n	(%)	
In situ	203	138	(27.1)*	65	(18.6)	
T1	369	250	(49.0)*	119	(34.1)	
T2	112	62	(12.1)	50	(14.3)	
Т3	81	28	(5.5)	53	(15.2)*	
T4	94	32	(6.3)	62	(17.8)*	
Total**	859	510	(100.0)	349	(100.0)	

Note: Chi-square test - p-value <0.001; \* = adjusted residuals analysis; \*\* = missing data

Table 4: Adjusted logistic regression according to histological level and age					
Variables	OR	95%CI	p-value*		
Age					
< 60 years	1.00				
≥60 years	1.64	1.22-2.20	0.001		
Histological level					
In situ	1.00				
T1	1.05	0.72-1.52	0.800		
T2	1.75	1.08-2.82	0.022		
T3	3.79	2.19-6.56	< 0.001		
T4	3.80	2.25-6.41	< 0.001		

Notes: OR = odds ratio; 95%CI = 95% confidence interval; \* = p-value, logistic regression

er than Brazil's national average, along with high socioeconomic indices such as mean *per capita* income, urbanization rate (96.6%), human development index (HDI) (0.809), and schooling, suggest that this population has a higher living standard than the Brazilian national average, and that it consists mostly of wage-earning urban workers with formal employment contracts and labor benefits.<sup>15-19</sup>

Such characteristics function as protective factors against the barriers to use of the SUS and provide greater ease in access to specialized treatments and tests.  $^{16,17}$ 

Our study design did not allow identifying the reason why the majority of patients diagnosed with CM were diagnosed in private practice. Patients' difficulties in using the public healthcare system (SUS) exclusively are known to potentially cause delays in diagnosis and treatment.<sup>17</sup> Patients thus trek through the health system, mixing different types of care, public and private, to the extent that they encounter difficulties in access to specialized care. This situation may lead SUS patients to seek medical care and pathology tests by paying out of pocket, as if they were from the private system. The barriers to use of services in the SUS can result in delays in diagnosis, and thus thicker CMs and even death. Another possible explanation for the predominance of patients from the private system in our series is underreporting of cases among SUS patients, as observed by Hoey et al. (2007) in patients with non-melanoma skin cancer (NMSC), among whom those from lower socioeconomic groups were less likely to seek medical care.20

In our series, patients in private care had more *in situ* and Breslow T1 cutaneous melanomas, while SUS patients had more T3 and T4 tumors. The comparison of median Breslow in the two groups showed the same result. Various studies have shown a positive correlation between thicker tumors at diagnosis and worse socioeconomic status. <sup>21-23</sup> Haenssle *et al.* (2015) found that higher educational level was a protective factor against higher-grade CM and that the delay in diagnosis and greater tumor thickness occurred in older patients, men, and those with less schooling. <sup>22</sup> Mayer *et al.* (2014) observed a higher risk of thick tumors (>2mm) in low-income patients. <sup>23</sup> Population-based studies have called attention to the fact that although the incidence of CM is significantly higher in patients with high income and high educational level, the prognosis is worse in patients with low socioeconomic status, low schooling, and un-

employment.<sup>20, 22</sup> It is worrisome that the longitudinal trend in the characteristics of patients and CMs in recent years has shown a higher increase in the incidence of Breslow T4 tumors in individuals with lower socioeconomic status and those over 65 years of age.<sup>23,24</sup>

Among the patients seen in the public Unified Health System (SUS), there was a predominance of the nodular melanoma histological type and elderly patients, while superficial spreading melanoma histological type and younger age were more common in private patients. A study in the United States also found an association between low socioeconomic status and nodular melanoma subtype and Breslow T4. Nodular melanoma accounted for only 14% of all the CMs in the study population, but it included a high proportion (37%) of fatal cases.<sup>23</sup>

It is worrisome that lower-income patients, who are mostly those treated exclusively in the public health system, have a diagnosis of more advanced stages of cutaneous melanoma and with worse prognosis. This reveals the inequality in Brazil's national health system, which formally recommends equal access to healthcare but is unable to meet this promise, thus excluding the more socially vulnerable population from timely access.<sup>22</sup>

The difficulty in access to specialists also contributes to worse prognosis. Studies have shown that dermatologists make the diagnosis earlier than physicians from other specialties, <sup>20,22-25</sup> and that their patients survive longer. <sup>22</sup> Residing in an area with a high density of dermatologists is independently associated with a diagnosis of thin CM. <sup>24</sup> Residing in an area with a low density of dermatologists and having a CM diagnosed by a non-dermatologist were independently associated with increased mortality from CM. <sup>8</sup> Patients that consulted a dermatologist were also more likely to perform self-examination of the skin. <sup>8</sup> Thus, studies show that dermatologists, as expected due to their training in the assessment of skin lesions, with or without dermoscopy, are more likely to diagnose and adequately manage pigmented lesions, when compared to all other medical specialists.

Participation in specialized screening programs in dermoscopy was the single strongest factor associated with diagnosis of thin CMs. Patients enrolled in these programs usually have an increased risk of CM, belong to younger age brackets at the time of diagnosis, and have more schooling.<sup>22</sup>

Our findings on cutaneous melanoma and socioeconomic factors require further elucidation in order to improve early detection and eliminate the socioeconomic disparities in access to diagnosis and treatment. The creation and improvement of compulsory reporting systems for cancers can help form a reliable population-based registry, crucially important for orienting interventions targeted at the population's behavioral characteristics and changes in public health policies.

Finally, early diagnosis of cutaneous melanoma reduces treatment costs, generating considerable savings for both the public healthcare system (SUS) and the private system. The funds resulting from such savings could be applied to programs for prevention and early diagnosis of skin cancer, as well as research on new techniques and drugs to treat these patients.<sup>26,27</sup>

The study's main limitation was its retrospective design using a secondary database. Our data may be underestimated since

some pathology specimens were sent from private offices to laboratories located outside the municipality, and some local residents may have gone elsewhere for diagnosis and treatment. In addition, some patients diagnosed with CM may have died. The data may also be overestimated because users of the public healthcare system (SUS) residing in other cities may have claimed the address of relatives or friends in order to gain access to medical services in Joinville, which is known as a reference in quality healthcare. However, given the size of the current study's case series, we believe that any possible bias was minimal and did not interfere greatly in the findings.

The exact mechanisms that explain our results are still not clear, but the difficulties faced by Brazil's public healthcare system (organizational, operational, and structural), along with the population's limited knowledge of cutaneous melanoma, less willingness to participate in screening programs, and other lifestyle factors not

analyzed in the current study may have contributed to late diagnosis in the group of patients seen in the public health system. More knowledge is needed on the role of these various sociodemographic, clinical, and behavioral factors, along with greater Breslow depth, in relation to late diagnosis of cutaneous melanoma both by healthcare providers and patients.

#### CONCLUSION

We found an independent association between histological level as verified by the Breslow index and the type of healthcare system coverage (p<0.001). Patients from the private healthcare system had thin cutaneous melanomas ( $in\ situ$  and Breslow T1), while patients seen exclusively in the public healthcare system (SUS) had thick cutaneous melanomas (Breslow T3 and T4).  $\square$ 

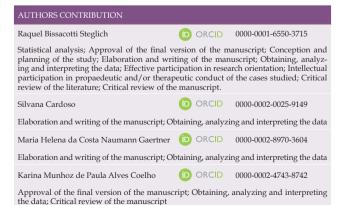
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