Burning mouth syndrome and associated factors: a retrospective study of 736 patients

Síndrome de boca ardiente y factores asociados: estudio retrospectivo de 736 pacientes

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Abstract

Background and objective. Burning mouth syndrome (BMS) can be defined as burning pain or dysesthesia on the tongue and/or other sites of the oral mucosa without a causative identifiable lesion. The discomfort is usually of daily recurrence, with a higher incidence among people aged 50 to 60 years, affecting mostly the female gender and diminishing their quality of life. The aim of this study was to evaluate the association between several pathogenic factors and burning mouth syndrome. Patients and methods. 736 medical records of patients diagnosed of burning mouth syndrome (BMS) and 132 clinical histories for the control group were studied retrospectively. The protocol included: gender, age, type of oral discomfort and other variables. **Results.** Analysis of the association between related factors and BMS diagnosis revealed that only 3 factors showed a statistically significant association (p = 0.0001): triggers (emotional factors, dental treatments), parafunctional habits, and oral hygiene. There was no statistically significant association between gender (p = 0.440), substance abuse (p = 0.089), systemic pathology (p = 0.099) and BMS. *Conclusions*. Parafunctional habits like bruxism and abnormal movements of tongue and lips can explain the BMS main symptomatology. Psychological aspects and systemic factors should be always considered. As a multifactorial disorder, the treatment of BMS should be executed in a holistic way.

Keywords: Anxiety; Bruxism; Burning mouth; Depression; Pain; Syndrome

Resumen

Antecedentes y objetivo. El síndrome de boca ardiente puede definirse como ardor o disestesia en la lengua y/u otras áreas de la mucosa oral, en ausencia de lesiones que puedan justificarlo. Estas molestias suelen recurrir diariamente, con mayor incidencia en pacientes de edades entre 50 y 60 años; predominan en el sexo femenino y provocan un deterioro de la calidad de vida. El objetivo de este estudio fue evaluar la asociación entre diversos factores patogénicos y el síndrome de boca ardiente. Pacientes y métodos. Se estudiaron de forma retrospectiva 736 historias clínicas de pacientes diagnosticados con síndrome de boca ardiente (SBA) y 132 historias clínicas de pacientes control. El protocolo incluyó: género, edad, tipo de molestia oral y otras variables. Resultados. El análisis de la asociación entre diversos factores y el diagnóstico de SBA mostró significación estadística (p = 0.0001) en solo 3 de ellos: factores desencadenantes (de orden emocional, tratamientos odontológicos), hábitos parafuncionales e higiene oral. No se encontraron diferencias significativas entre el género (p = 0.440), abuso de sustancias (p = 0.089), patología sistémica (p = 0.099) y SBA. Conclusiones. Los hábitos parafuncionales como el bruxismo y los movimientos anormales de la lengua y labios pueden explicar la sintomatología del SBA. Hay que tener en cuenta siempre los aspectos psicológicos y los factores sistémicos. Como alteración de carácter multifactorial que es, el tratamiento del SBA debe contemplarse de manera holística.

Palabras claves: Ansiedad; Bruxismo; Ardor bucal; Depresión; Dolor; Síndrome.

Introduction

Burning mouth syndrome (BMS) is mostly expressed as a "burning or stinging sensation on the oral mucosa", "oral dysesthesia", "burning pain on the tongue" and so on. Without a causative identifiable lesion, the discomfort is usually of daily recurrence, although periods of no pain during the day are reported. It occurs mostly in females, affecting their quality of life^{1–4}.

The International Association for the Study of Pain (IASP) defines BMS as burning pain or dysesthesia on the tongue and/or other sites of the oral mucosa, without any clinical or laboratory data. This syndrome has been classified in primary and secondary. In the primary form (essential or idiopathic), the burning sensation is not accompanied by clinical or analytical alterations; in the secondary form, lesions are present, with laboratory abnormalities, systemic diseases, psychological alterations, or consumption of specific medicines⁵. Lamey and Lewis (1989) proposed three clinical types based on the diurnal variation of the symptoms. The discomfort tends to be chronic, but spontaneous periods of remission are also common³. The typical patient with BMS is a menopausal woman with systemic diseases, burning mouth sensation on the tongue, taste alterations and dry mouth, without clinical lesion⁷.

Prevalence ranges between 0.7% and 5.1% in general population^{1,8}, and 33% in risk groups¹. Incidence is higher among people aged 50 to 60 years, with a ratio women/men between 3:1⁸ and 9:1¹. This difference has been related to biological, psychological, and social factors^{9–12}. The psychological component in BMS is supported by studies that report higher levels of depression and anxiety in patients with BMS compared to control groups¹². In a study performed on 30 patients with BMS a significantly association was found between taste alteration (metallic or

acidic taste) and depression¹³. Other studies showed that depression and anxiety were present in 50% of cases with BMS^{1,14} and 67.1% of patients with this syndrome were found to suffer poor sleep quality¹⁰. An association of openness personality trait with stress-related salivary biomarkers has been related to BMS¹⁵. In recent years, new evidence seems to link BMS with peripheral and neuropathic disturbances^{11,16}. Menopausal hormone reduction has also been linked to BMS^{2,3,9}. Among the reported systemic factors associated with this syndrome are: diabetes mellitus¹⁷, gastrointestinal reflux¹⁸, nutritional deficiencies¹⁷, hormonal changes⁹, and some adverse effects of drugs¹⁷. Local factors and clinical conditions found in these patients include geographic tongue, candidiasis, parafunctional habits, hyposalivation¹⁷, as well as changes in the saliva composition, contact allergies¹⁹ and dental treatments (prosthesis, implants, teeth extractions)²⁰.

Management of BMS should be focused against the etiopathogenic factors, based on preventing the causes of oral irritation. However, in some patients etiological factors are not easy to identify, in that case BMS is considered to be idiopathic^{2,11,16}. Several patients report improvement with cold beverages, soft food and/or recreational activities⁴. The use of pharmacological treatment, particularly topic clonazepam can also help to ameliorate the symptoms.²¹ Psychological support does usually help¹².

Differential diagnosis of BMS includes: candidiasis, oral lichen planus, geographic tongue, contact allergic stomatitis, and xerostomia. Systemic diseases such as Sjögren syndrome and other forms of dry mouth²², vitamin and mineral deficiencies²³, uncontrolled diabetes and hematological diseases²⁴, should also be ruled out. If any of these are the causal factors, their

treatment should resolve the burning sensation. For idiopathic cases, several treatments have been proposed to improve BMS, with variable results^{2,14,21,25}.

Parafunctional habits like bruxism and abnormal movements of tongue and lips seem to play a significant role in the pathogenesis of BMS^{7,25–27}. Since the etiology of BMS is multifactorial, the main objective of this study was to evaluate the association between several pathogenic factors and the semiology of BMS.

Patients and methods

736 medical records of patients diagnosed of BMS and 132 clinical histories for the control group were studied retrospectively. Diagnosis was established by three experts in oral medicine. The protocol included: gender, age, kind of pain or dysesthesia and time span of the burning sensation. Other variables considered were: location and intensity of oral discomfort, triggers (like recent dental treatment or emotional factors), parafunctional habits (like bruxism, *morsicatio* and other repetitive movements of tongue, lips or other oral structures), systemic pathology, oral hygiene, substance abuse (alcohol, tobacco, etc.), and type of treatment (Table 1). All patients diagnosed by BMS were visited during the last 25 years. The application form was approved by the Ethical Committee of the Dental Hospital, University of Barcelona.

Ethical aspects

To maintain the patient confidentiality, recommendations of the Spanish Organic Law 15/1999 of December 13 on the Protection of Personal Data were followed. All participants were identified by a code only known by the researchers.

Inclusion criteria

Patients of any gender and age suffering from burning mouth sensation and diagnosed of any type of BMS were included. Patients with burning mouth sensation attributable to other causes (oral candidiasis, geographic tongue, oral lichen planus) were excluded.

Control group

The BMS group of patients was compared with a control group consisting of subjects who did not experience a burning sensation or discomfort in the mouth. These subjects were unselected consecutive patients attending the dental clinic during the last year. The gender and age distributions of the control group were similar to those of the BMS group.

Statistical analysis

The data were analyzed with the SPSS Statistics software package (ver. 20.0, IBM), and $p \le 0.05$ was required for statistical significance.

The normality of the distribution of continuous variables (age and time of duration) was tested by means of the Kolmogorov-Smirnov test. To describe the incidence of the discrete variables, the percentages for the combined sample and for gender-segregated samples were computed. To investigate the association between diagnosed BMS and each factor considered as variable, a multinomial logistic regression model was used, appropriate for dichotomous factors (presence/absence). In addition to the multiple regression analysis conducted with all the factors combined, the associations between each factor and BMS were analyzed separately by means of the chi-square (χ^2) test.

Results

The total sample included 736 patients with BMS and 132 control patients (Table 2). The ratio of women to men in the BMS group was 6:1, similar to the control group. The average age of the patients in the BMS group were 60.63 years for women and 59.07 years for men; in the control group, the corresponding ages were 55.11 and 55.48 years. Age was normally distributed in both groups (p = 0.608 for the control group and p = 0.168 for the BMS group). These results indicate that both samples were random and that there was no bias about age distribution. In contrast, the distribution of data for duration of BMS deviated substantially from normal, because of the variation between the onset of the symptoms and the diagnosis (Table 2).

Analysis of the factors associated with BMS

Analysis of the associations between the factors and the diagnosis of BMS (Table 3, Analysis 1) revealed three significant associations (p = 0.0001 in all three cases): triggers (TRIG), parafunctional habits (PARH), and oral hygiene (HYG). There was no association between gender and BMS (p = 0.440), and therefore no differential affectation was observed by gender. Neither substance abuse (SABU, p = 0.089) nor systemic pathology (SYSTP, p = 0.099) were significant factors in the development of BMS, although the p values approached statistical significance (p < 0.10).

To assess the specific weight of each factor, a second regression analysis (without the not significant factors of the previous one) was carried out. Note, however, that substance abuse and systemic pathology were included in the second analysis, because p was smaller than 0.10 for these factors. In this second analysis, triggers, parafunctional habits, and oral hygiene maintained their significance, and in addition, substance abuse showed a significant association with BMS (p)

= 0.012). In contrast, systemic pathology remained not significant, and in fact, the p value increased to 0.131 (Table 3, Analysis 2).

The results shown in Table 3 suggest an association between BMS and triggers, parafunctional habits, hygiene, and substance abuse (which is related to hygiene, usual less oral hygiene in smokers).

Finally, associations between factors (Table 4) were analyzed by means of contingency tables of differences, and the results showed that diagnosis of BMS was clearly associated with triggers, parafunctional habits, and hygiene, as well as with systemic pathology. This result suggests that systemic pathologies may have significant weight in BMS, despite the fact that the initial logistic regression analysis did not show a significant association (p > 0.05). In this analysis, substance abuse was not significantly associated with the definitive diagnosis of BMS. With respect to the association between the other factors, all of them proved to be significantly associated ($p \le 0.035$). This result suggests that BMS is multifactorial.

Discussion

The results suggest that BMS is associated with various factors and is more frequent among females than among males (6:1), in agreement with previous reports^{1,25,28}. Hormonal status may be involved in the greater prevalence in women, perhaps by modulating other factors in some way². The average age of patients in the BMS group was 60.4 years, (range, 22–97 years). Middle-aged and elderly patients tend to have multiple systemic pathologies and consequently tend to be subjected to several treatments, most of them causing xerostomia. Habitual smoking (tobacco, marijuana) also lead to local dryness and worsen the clinical condition of the

syndrome⁷. Dry mouth in combination with some triggers (psychological–anxiety, depression; dental treatments; poor oral hygiene; substance abuse; etc.)¹², tend to promote some parafunctional habits like bruxism, repetitive swallowing, and diverse movements of tongue and lips⁷. Patients with BMS exhibit significantly more symptoms of depression and anxiety compared with healthy subjects¹⁰; this may explain the parafunctional habits and BMS association found in our study.

The variables "parafunctional habits", "triggers", "poor oral hygiene", and, to a lesser extent, "substance abuse" and "systemic pathology" were clearly associated with BMS. Parafunctional habits have repercussions for the entire orofacial area. Bruxism can lead to dental and periodontal hypersensitivity. Corsalini et al. found in their study of 44 patients with BMS that 72.2% showed parafunctional habits; especially night bruxism was observed in twenty-eight cases, clenching in ten cases, biting of lips and cheeks in six cases²⁹. *Morsicatio* of the cheek or lip's mucosa, tongue thrusting against teeth or prosthesis and other similar habits, can sometimes be performed continuously or intermittently generating local discomfort that involves teeth, oral mucosa, joints, and muscles. Parafunctional habits (related to tongue and lips) often coincide with poor oral hygiene (presence of plaque or calculus). The repetitive movement of tongue and lips causes tissue irritation; this can alter the perception of taste¹³, sometimes transmitted to the brain as distorted information of the peripheral stimulus (burning sensation)⁷.

Probably, the stress conditions associated to BMS could explain in these patients the onset of parafunctional habits²⁹. Trigger of symptoms can often be of emotional kind (serious illness, death of a relative, problems at work), and they are also frequently associated to a recent dental treatment. The patient usually links the trigger to the symptoms. In a study performed on 75

patients with BMS, the most frequent trigger reported was stressful life events (45%) and an antecedent dental procedure was present in 20% of the cases³⁰.

In conclusion, the etiology of BMS is not fully known, making its diagnosis a challenging process; several factors are associated with this pathology, what does justify a multifactorial approach and treatment. BMS management must always include the psychological aspects; many times the patient's externalization of the complaint can help them feel reassured. Systemic associated factors like xerostomia, dehydration, or underlying nutritional deficiencies should always be treated. Local factors like oral hygiene, elimination of local irritants (rinses, toothpastes), and parafunctional habits should be controlled (a lower splint may be helpful). Sometimes, the use of pharmacological treatment such as topical clonazepam can also help to ameliorate the symptoms.

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Table 1. Analyzed variables and factors considered for their association with burning mouth syndrome.

Variable or factor	Description of discrete values	
Reason for	BURNING MOUTH (or its variations, e.g.,	
consultation	stinging, sour taste, sandy feeling)	
Consultation	NO BURNING	
Type of discomfort	ONLY BURNING	
Type of discomfort	BURNING + OTHER (e.g., dysgeusia, dryness)	
Location	TONGUE + OTHER (e.g., lips, palate, gums)	
Location	NOT ON TONGUE	
Pain intensity (analog	≤5	
scale)	>5	
	PRESENT (emotional triggers, recent dental	
Triggers	treatments)	
	ABSENT	
Parafunctional habits	PRESENT (e.g., dental, lingual, muscular, mucosa)	
r ararunctional naoits	ABSENT	
Ct	PRESENT (associated with medication)	
Systemic pathology	ABSENT (or does not take medication)	
	GOOD	
Oral hygiene	AVERAGE OR POOR (presence of bacterial	
	plaque, calculus)	
	GOOD (balanced diet, good hydration, i.e., >2 L of	
Dietary habits	liquid/day)	
	BAD (inappropriate diet, poor hydration)	
Substance abuse	PRESENT (e.g., tobacco, alcohol, marijuana)	
Substance abuse	ABSENT	
	PRESENT (maladjusted prosthesis, sharp teeth or	
Local factors	filling edges, etc.)	
	ABSENT	
Existing treatment	CONSERVATIVE (hydration, saliva substitutes,	
	hygiene, balanced diet)	
	MEDICINE (e.g., clonazepam, antidepressants,	
	sedatives)	
Definitive diagnosis	BMS	
	no BMS or OTHER pathologies	

Table 2. Distribution by gender and age of the BMS and control groups.

	BMS (r	a = 736)	Control $(n = 132)$		
Age (years)	x = 60.41 SD = 13.38 range = 22–97 z = 1.112 p = 0.168	Women, $n = 630$ x = 60.63 SD = 13.25 z = 0.961 p = 0.314 Men, $n = 106$ x = 59.07 SD = 14.14 z = 0.632 p = 0.819	x = 55.18 SD = 14.89 range = 24–91 z = 0.762 p = 0.608	Women, $N = 107$ x = 55.11 SD = 14.64 z = 0.628 p = 0.826 Men, $N = 25$ x = 55.48 SD = 16.22 z = 0.702 p = 0.709	
Time of evolution (months)	x = 21.19 SD = 39.77 range = 0.25–600 $z = 8.186$ $p = 0.000$	Women, $N = 630$ x = 21.17 SD = 39.91 z = 7.577 p = 0.000 Men, $N = 106$ x = 21.31 SD = 39.06 z = 3.214 p = 0.000			

Abbreviations: x, mean value; SD, standard deviation; z, Kolmogorov-Smirnov statistic for normality of the analyzed variable; p, significance of the Kolmogorov-Smirnov z test (if p < 0.05, the distribution of the variable was considered to deviate significantly from normal). Significant p values are shown in bold.

Table 3. Logistic regression analyses 1 and 2 of associations between main factors involved in BMS development.

	Analysis 1			Analysis 2		
	-2 log v	χ^2	p	-2 log v	χ^2	p
Intercept	245.403			133.368		
GEN	245.998	0.595	0.440			
TRIG	266.507	21.104	0.000	157.254	23.886	0.000
PARH	269.971	24.568	0.000	167.211	33.843	0.000
SYSTP	248.124	2.721	0.099	135.650	2.282	0.131
HYG	269.679	24.275	0.000	157.659	24.291	0.000
DIET	245.622	0.219	0.640			
SABU	248.302	2.899	0.089	139.721	6.353	0.012
LOCF	246.04	0.637	0.425			

Abbreviations: GEN, gender; TRIG, triggers; PARH, parafunctional habits; SYSTP, systemic pathology; HYG, oral hygiene; DIET, dietary factors; SABU, substance abuse; LOCF, local factors. $-2 \log v =$ statistic of the reliability (v) of the logistic regressions; $\chi^2 =$ chi-square statistic for the association between the dependent variable (BMS) and the factors considered. Analysis 1 included all the analyzed factors, whereas analysis 2 included only the factors that were significant (or approached significance) in analysis 1 (p < 0.100). Significant p values are shown in bold.

Table 4. χ^2 and p values for associations between BMS and the factors considered.

	BMS
TRIG	$\chi^2 = 57.369$
TRIO	p = 0.000
PARH	$\chi^2 = 34.841$
ГАКП	p = 0.000
HYG	$\chi^2 = 40.249$
1110	p = 0.000
SABU	$\chi^2 = 0.088$
SABU	p = 0.993
SYSTP	$\chi^2 = 8.764$
	p = 0.033

Abbreviations: BMS, positive definitive diagnosis of Burning Mouth Syndrome; TRIG, triggers; PARH, parafunctional habits; HYG, oral hygiene; SABU, substance abuse; SYSTP, systemic pathology.