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ASSOCIATION BETWEEN CAFFEINE, ANXIETY AND THE OCCURRENCE OF APHTHOUS STOMATITIS IN THE ARMENIAN ETHNICITY

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Abstract

Introduction: Aphthous stomatitis is a recurring oral ulceration condition that affects a significant proportion of the population. While several factors have been proposed to contribute to its occurrence, the link between caffeine consumption, anxiety levels, and the development of aphthous stomatitis remains unknown. This survey-based study aimed to understand the association between coffee, anxiety, and the development of aphthous stomatitis in this specific demographic, which can provide useful insights for medical management as well as preventive strategies.

Material and methods: A structured online questionnaire was distributed via various social media platforms targeting a sample of the Armenian population. The questionnaire collected information on participants' demographics, coffee use patterns, anxiety levels as measured by perceived stress scales, and the occurrence of aphthous stomatitis. Statistical analyses, including Kendall's tau, were carried out to investigate the relationships.

Results: The results revealed a significant negative association between caffeine consumption and the occurrence of aphthous stomatitis among Armenian individuals Kendall's tau-b (τ b) correlation - 240* (p<0.05). Participants who reported regular caffeine consumption, such as coffee, tea, and energy drinks, had a lesser prevalence of aphthous stomatitis than non-consumers. Furthermore, Armenians in the survey showed that the majority (83.5%) had a low to moderate stress level.

Conclusion: The data we collected in our research indicate that the prevalence of Recurrent Aphthous Stomatitis in the Armenian ethnicity was infrequent; over half of the participants didn't report Recurrent Aphthous Stomatitis in the past 12 months. Which may correlate to their caffeine consumption habits, which can lead to stress reduction.

Keywords: aphthous stomatitis, caffeine, anxiety, stress, Armenian ethnicity

INTRODUCTION

Recurrent Aphthous Stomatitis (RAS) belongs to the group of chronic inflammatory diseases of the oral cavity, the characteristic feature of the disease is single or multiple shallow round ulcerations usually occurring in non-keratinized areas like the lips, ventral surface of the tongue, floor of the mouth, buccal mucosa and soft palate. The incidence of RAS ranges from 5% to 50% depending on the ethnic or socioeconomic group [*Chattopadhyay A, Chatterjee S, 2007; Akintoye S, Greenberg M, 2014*].

Despite all the efforts, the exact aetiology of RAS is still ambiguous, and both genetic and environmental factors are thought to be implicated. the precipitating factors include stress, physical or chemical trauma, allergies, infection, drugs or nutritional deficiencies. Stress is believed to be a

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Maryam H. Alwan, M.Sc. College of Dentistry, University of Baghdad Babalmoadhum 1417, Baghdad Tel.: (+964) 26-28-819 E-mail: maryam.h@codental.uobaghdad.edu.iq major risk factor for RAS due to its effect on salivary cortisol levels, para-functional habits that lead to self-inflicted trauma and alterations to the immune system [*Albanidou F et al.*, 2008].

Anxiety is known as the body's response to stress, and there is an ongoing debate over the effects that caffeine consumption has on anxiety since caffeine is known as the most widely consumed psychostimulant substance in the world [Lara D, 2010; Al-Bazaz N, Radhi N, 2021].

Many cited evidence has shown different results, some stated that there was no correlation between caffeine and stress or anxiety [Hall S et al., 2015], Other research stated that very high doses can increase stress levels while moderate to low doses helped in reducing stress levels [McLellan T et al., 2016]. Since stress is a multifactorial condition, it's better to research specific individuals with similar lifestyles to accurately measure their stress levels and caffeine habits, therefore in this paper, we will focus on the Armenian ethnicity, where genetics and cultural customs are shared between the subjects.

Coffee consumption is a popular cultural habit in Armenia, and some studies suggest that social factors and cultural norms may play a role in the high prevalence of coffee consumption among Armenians. For example, a study by Grigoryan et al. (2018) found that social interactions and traditions associated with drinking tea and coffee were important factors in the dietary patterns of the Armenian population. Daily consumption of coffee in various age groups is a practice that Armenians associate with their routine life, coffee is a must with company, celebrations and for a peaceful afternoon rest, therefore it is expected that the abovementioned effects of chronic caffeinating on the body to be vibrant in the Armenian ethnicity [Aleksanyan A et al., 2020; Pipoyan D et al., 2023].

MATERIAL AND METHODS

Study design and sample preparation: The self-administered questionnaire was conducted using Google Forms and distributed to the participants via various social media platforms. The survey mainly targeted a sample of the Armenian population. Two online forms were created with identical questions but in different languages; the first form was in English and the second in Armenian, to accommodate the language preferences of the participants and make sure the questions were clear and understood by everyone. They were asked to participate voluntarily and comply with Baghdad University College of Dentistry's experi-

mental ethical guidelines. All the participants were fully informed about the aims of the study and the confidentiality of the data, they were also assured that the data would be used only for research purposes and that no name or IP address was being recorded to protect their privacy and the confidentiality of their personal information. Table 1 summarises the elements of the questionnaire, which consists of three sections: Caffeine Consumption Assessment (Table 1a), Recurrent aphthous stomatitis frequency assessment (Table 1b) and Perceived Stress Scale (Table 1c).

Statistical analysis: Statistical analysis was conducted using the SPSS software, version 26. Descriptive statistics for frequency, mean, and standard deviation. Student T-test was used for comparisons between two independent groups. In addition, Kendall's tau-b (τ b) correlation coefficient was applied to find the strength and direction of association that exists between the variables. The p<0.05 value was considered statistically significant.

Correlation between Variable: First and foremost, the amount of coffee consumed by each individual was calculated by multiplying the number of cups consumed by the frequency of consumption per day. Subsequently, to find the correlation between the two variables, the number of caffeinated drinks taken per day in ml and the occurrence of aphthous stomatitis, Kendall's tau-b (τ b) correlation coefficient was applied.

Results

The questionnaire was found to be reliable; Cronbach's alpha coefficients for variables were 0.715 which is considered to have satisfactory internal consistency. A total of 97 answers were collected, with female participants accounting for 78.4% of the total and male participants accounting for 21.6%. The majority are in their twenties and thirties. 55.7% were university students or graduates, while 24.7% had postgraduate degrees, and 19.6% were high school students or lower. Table 2 summarises the demographic characteristics of the participants.

Recurrent aphthous stomatitis Questionnaire response: The study's main finding was that 70% of the participants had an ulcer six months to a year ago. 52.6% had only one frequency, 53.6% had them for 3-5 days, and 73.2% had minor or no pain. The response to the RAS questionnaire is summarized in table 3.

The Perceived Stress Scale response: *Evaluation Results of the Perceived Stress Scale:* The

_	<i>Tabl</i> stionnaire in English an		
Armenia	Armenian language		
Caffeine Consur	nption Assessment		
Questions	Select the answer and indicate		
	Coffee		
	Tea		
Sources of caffeine	Caffeinated soft drinks		
Sources of calleline	Energy Drinks		
	Chocolate		
	Others		
	Once a day		
	2-3 times daily		
How often do you consume	e4-5 times daily		
Caffeinated beverages?	More than 5 daily		
	1-3 times weekly		
	4-6 times weekly		
What time of the day	Morning		
do you usually consume	Afternoon		
caffeine?	Evening		
Wilson de sees ander te	Before eating		
When do you prefer to have caffeine?	While eating		
	After eating		
	13-17		
At what age did you start	8-12		
taking caffeine ?	Under 8		
	Over 18		
	Alert, energized and		
Caffeine makes me	less sleepy		
	Think clearly, pay more		
	attention and sharpen		
	my memory		
	Relax, calm down, and		
	improves my mood		
	$\frac{S(240 ml)}{M(260 ml)}$		
Cup Size	M (360 <i>ml</i>)	_	
1	L (480 ml)		
	XL (600 ml)		

TABLE 1b

The element of the questionnaire in English and Armenian language

Recurrent aphthous stomatitis frequency		
ass	essment	
Questions	Select the answer and indicate	
Did you experience any	Yes	
type of mouth ulcer before?	No	\checkmark
	Currently have it	
When was the last time	One month ago	
you experienced an ulcer ?	6 months ago	
	One year ago or more	
	1 time	
Frequency of Ulcer ?	2-3 times	
	4 or more	
	None	
	0-2	
How long do they last? (Days)	3-5	
(Days)	6 or more	
	No pain	
II	Slight pain	
How painful is it ?	Moderate pain	
	Severe pain	
	No treatment	
Any treatment taken ?	Topical gel or vitamins	
	Home remedies	
	Salty	
Do you regularly eat food that is	Acidic	
	Spicy	
	None	

TABLE 1c

The element of the questionnaire in English and Armenian language

Perceived Stress Scale	
F In the left column for each question., we use a scale ranging from $(0-5)$ where $0 =$ never, $1 =$ almost, never, 2	=
sometimes, 3 = fairly often and 4 = very often	
1. In the last month, how often have you been upset because of something that happened unexpectedly?	2
2. In the last month, how often have you felt that you were unable to control the important things in your life?	4
3. In the last month, how often have you felt nervous and stressed?	1
4. In the last month, how often have you felt confident about your ability to handle your personal problems?	0
5. In the last month, how often have you felt that things were going your way?	5
6. In the last month, how often have you found that you could not cope with all the things that you had to do?	2
7. In the last month, how often have you been able to control irritations in your life?	2
8. In the last month, how often have you felt that you were on top of things?	2
9. In the last month, how often have you been angered because of things that happened that were outside of your control?	3
10. In the last month, how often have you felt difficulties were piling up so high that you could not overcome them?	4

stress levels of the 97 participants in the survey showed that the majority (83.5%) had a low to moderate stress level, which was distributed as low, moderate, or high, accounting for 24.74%, 58.76%, and 16.49%, respectively [*Cohen S et al., 1983; Siqueira Reis R et al., 2010*].

Caffeine Consumption Questionnaire and Response: The results of this section of the questionnaire revealed that 69.2% of respondents consume two or more sources of caffeine, 49.5% con-

		IABLE 2
Demographic characteristics of participants		
Variable	Category	n (%)
Gender -	Male	21 (21.6)
	Female	76 (78.4)
Age	<20	5 (5.2)
	20-40	77 (79.4)
	>40	15 (15.5)
Education -	University student or graduate	54 (55.7)
	Postgraduate (Master's, Ph.D.)	24 (24.7)
	High school or less	19 (19.6)

TABLE 2

TABLE 3

Recurrent aphthous stomatitis questionnaire and

response		
Questions	Answers	n (%)
Did You Experience	Yes	80 (82.5)
Any Type of Mouth Ulcer Before?	No	17 (17.5)
	Currently Have It	5 (5.2)
When Was the Last	1 Month Ago,	23 (23.7)
Time You Experienced	6 Months Ago,	17 (17.5)
an Ulcer	1 Year Ago, & More	52 (53.6)
	None	17 (17.5)
F 0.411 0	Once	51 (52.6)
Frequency Of Ulcer?	2-3	23 (23.7)
	4 And more	6 (6.2)
	0-2	31 (32)
How Long Do They	3-5	52 (53.6)
Last? (Days)	6 And more	14 (14.4)
	No Pain	31 (32)
How Painful Is It?	Slight Pain	40 (41.2)
now Painful Is It?	Moderate Pain	24 (24.7)
	Severe Pain	2 (2.1)
	No Treatment	66 (68)
Any Treatment Taken?	Topical Gel or	19 (19.6)
	Vitamins	19 (19.0)
	Home Remedies	12 (12.4)
Do You Regularly Eat Food That Is	none	24 (24.7)
	Salty	21 (21.6)
	Acidic	7 (7.2)
	Spicy	17 (17.5)
	Acidic and salty	20 (20.6)
	spicy and salty	8 (8.2)

sume it more than three times per day, and 51.5% begin using caffeine before the age of eighteen (Table 4).

Moreover, when asked how caffeine makes them feel, 51.5 percent said to relax, calm down, and improve their mood, 38.1 percent said alert, energized, and less sleepy, and 10.3 percent said improves their mood. Think clearly, pay more attention, and sharpen the memory (Figure).

Correlation between Variable: Kendall's tau-b (τ b) correlation coefficient was applied. The result is interpreted as follows: A correlation value of -.240* between two variables indicates that they have a significant and negative association. A negative correlation suggests that as the value of one-factor increases, the value of the other decreases (Table 5).

DISCUSSION

The impact of oral lesions such as RAS on the quality of life is considered very important as these lesions affect speech, nutrition and general comfort, and the incidence of this condition is found to be higher in individuals who experience high levels of

		TABLE 4
Caffeine consumption questionnaire response		
Questions	Answers	n (%)
	Coffee	18(18.5)
	Tea	8(8.2)
Sources of coffeine	Chocolate	3 (3.1)
Sources of caffeine	Energy Drinks	1 (1)
	two types	35(36.1)
	more than two types	32 (33.1)
	Once a day	28(28.9)
Π	2-3 times daily	43(44.3)
How often do you consume Caffeinated	4-5 times daily	5(5.2)
	More than 5 daily	2(2.1)
beverages?	1-3 times weekly	9 (9.3)
	4-6 times weekly	10 (10.3)
	Morning	28 (28.9)
	Afternoon	19 (19.6)
What time of the	Evening	15 (15.5)
day do you usually	Morning and	12 (12 4)
consume caffeine?	Evening	12 (12.4)
	Morning and	22 (22 7)
	Afternoon	23 (23.7)
Wilson de soon ander	Before eating	23 (23.7)
When do you prefer to have caffeine?	While eating	16 (16.5)
to have caffeine?	After eating	58 (59.8)
At what are did you	8-12	8 (8.2)
At what age did you start taking caffeine? (years)	13-17	46 (47.4)
	Over 18	39 (40.2)
	Under 8	4 (4.1)
Cup Size (ml)	240	41 (42.3)
	360	39 (40.2)
	480	16 (16.5)
	600	1 (1)

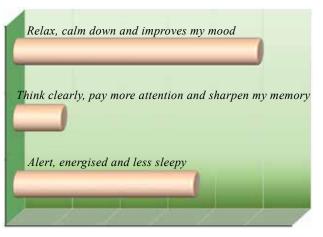


FIGURE. How the participant thinks caffeine makes them feel?

stress [*Thevara M et al., 2020*]. Caffeine being the most widely consumed psychostimulant substance has been reported to have beneficial effects on certain neurotransmitters such as dopamine and sero-tonin which in turn lead to mood improvements and feelings of wellbeing [*Singh K, 2016*].

In this study, questionnaires were collected from individuals of Armenian ethnicity to explore their caffeine habits, stress levels, and RAS history. And according to our data, it's been found that more than eighty percent of the participants had low to moderate stress levels, and fifty percent felt that coffee made them feel relaxed, calmed down, and improved their mood. These findings correlate with a previous study done by Haskell *et al.*, where they did a double-blind test to compare the effects of regular coffee, decaffeinated coffee, and placebo, it was found that regular coffee had an overall higher mood improvement and decreased mental fatigue [*Haskell R et al., 2018*].

Similar results were reported in previous literature. A study published in the Journal of Psychopharmacology noticed a reduction in stress levels in both

TABLE 5

Correlation between Variable. Kendall's tau-b (τb)

correlation			
	Kendall's tau_b	X (n=97)	Y (n=97)
Х	Correlation Coefficient	1.000	-0.240*
(n=97)	Sig. (2-tailed)	0.0	0.015
Y	Correlation Coefficient	-0.240*	1.000
(n=97)	Sig. (2-tailed)	0.015	0.0
Note: * Correlation is significant at the 0.05 level (2-tailed). Kendall's tau-b (τ b) correlation representation in blue cells. The white cells represent the maximum values of Kendall's tau-b (τ b) correlation, X -Experienced any type of mouth ulcer before?, Y-Amount of caffeinated taken per day in ml			

men and women who consume caffeine [*Smith A*, 2002; *Heckman M et al.*, 2010; *Socała K et al.*, 2020].

The data also suggest that more than half of the participants haven't had RAS in the last twelve months, this may be associated with the low-stress levels which agrees with the results of the study done by Albanidou et al. (2008), who found a positive association between RAS and the participant's anxiety levels. While the direct relationship between coffee consumption and aphthous stomatitis has not been thoroughly investigated, there is evidence that stress reduction through coffee consumption may indirectly contribute to a decrease in its prevalence. A study by Altenburg et al. (2008) concluded that chronic stress suppresses the immune system and can result in inflammatory disorders such as aphthous stomatitis. Coffee consumption may help modify the body's immunological and inflammatory responses, potentially decreasing the occurrence of aphthous stomatitis, by lowering stress levels [Altenburg A et al., 2008].

Furthermore, an interesting study published in the Biomedical and Pharmacology Journal found that Methylxanthines are a kind of naturally occurring chemical present in coffee, tea, chocolate, and other foods. The only substances in therapeutic use are naturally occurring theophylline (1,3-dimethylxanthine) and its ester derivative aminophylline. They have a similar molecular structure to caffeine. It also has anti-inflammatory qualities by reducing cytokine synthesis and the action of cytokines on leukocytes. According to several studies, pentoxifylline is useful in avoiding aphthous ulcers [*Sridhar T et al., 2015; Alwan M, ghani B, 2020; Bondarev A et al., 2022*].

Due to the vague aetiology, even in the present-day research and studies on RAS continues to be noteworthy to be able to understand the condition more.

Conclusion

While there is little direct research on the relationship between coffee consumption, stress reduction, and the prevalence of aphthous stomatitis, the current data shows a possible indirect association. The data we collected in our research found that the prevalence of RAS in the Armenian ethnicity was infrequent, over half of the participants didn't get RAS in the past 12 months. Which may correlate with their caffeine consumption habits, which can lead to stress reduction. However, caution should be taken when consuming caffeine, as harmful effects are potential with large doses.

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THE NEW ARMENIAN MEDICAL JOURNAL



Volume17 (2023). Issue 4



CONTENTS

4. AVAGYAN S.A., ZILFYAN A.V., MURADYAN A.A.

SELECTIVE ADMINISTRATION OF POLYAMINE-DEFICIENT AND POLYAMINE-FREE DIETS TO CANCER PATIENTS

17. Alshehri K., Morsi N., Mahsoon A.

THE EFFECT OF WORKPLACE BULLYING ON NURSES' MENTAL WELL-BEING IN SAUDI ARABIA

31. SADUAKAS A.Y., KURAKBAYEV K.K., MATKERIMOV A.ZH., TERGEUSSIZOV A.S., SAGATOV I.Y., SHAMSHIYEV A.S., ZHAKUBAYEV M.A., BAUBEKOV A.A., TAJIBAYEV T.K., KHANSHI MEAD, KOZHAMKUL A.ZH., MADADOV I.K.

THE BENEFITS OF DUPLEX SCANNING OF EXTRACRANIAL CAROTID PATHOLOGIES FOR RISK STRATIFICATION OF ISCHEMIC STROKE

36. Soleimantabar H., Sabouri S., Shirbandi K.

RISK OF CARDIAC ANOMALIES IN ABERRANT RIGHT SUBCLAVIAN ARTERY RELA-TIVE AORTIC ARCH ANOMALIES FOR PEDIATRICS: A CROSS-SECTIONAL STUDY

42. Abbaspour M., Hejazi Z.S., Namjoyan F., Azemi M.E.

FORMULATION OF VAGINAL CREAM CONTAINING EXTRACTS OF LINUM USITATISSIMUM, FOENICULUM VULGARE, AND SALVIA OFFICINALIS FOR THE TREATMENT OF ATROPHIC VAGINITIS IN POSTMENOPAUSAL

48. ZARGAR M., NAJAFIAN M., SHOJAEI K., MORADKHANI N.

AN INVESTIGATION INTO THE IMPACT OF CONTINUING OR TERMINATING PREGNANCY ON THE MATERNAL, FETAL AND DISEASE PROGRESSION OUTCOMES IN PREGNANT WOMEN WITH COVID-19

58. MARTUSEVICH A.K., FEDOTOVA A.S., SUROVEGINA A.V., NAZAROV V.V.

PLASMA BIOMEDICINE: MODERN STATE-OF-ART AND PERSPECTIVES IN REGENERATIVE MEDICINE

66. Martusevich A.K., Kovaleva L.K., Fedotova A.S., Stepanova E.A., Soloveva A.G. EXPERIMENTAL STUDY OF ERYTHROCYTE ENERGY METABOLISM UNDER INHALATIONS OF NITRIC OXIDE

71. BAKHTARI A., GAVANJI S.

IN-SILICO DOCKING ANALYSIS OF SELECTED FLAVONOIDS AND PROTECTIVE ANTIGEN

77. KHERADMAND P., SHAFEENIA R., BAGHERI S., ATTAR A.

CAUDAL TYPE HOMEOBOX 2 EXPRESSION AND PROGNOSTIC FACTORS IN PATIENTS WITH GASTRIC ADENOCARCINOMA

84. DARMADI D.

IMPROVEMENT OF ENZYME IMMUNODETECTION IN THE LABORATORY DIAGNOSIS OF HEPATITIS E VIRUS

98. TKHRUNI F.N., ISRAYELYAN A.L., KARAPETYAN K.J.*, BALABEKYAN TS.R., KHACHATRYAN L.M., KHACHATRYAN T.V.

COMPARATIVE ANTIMICROBIAL ACTIVITY OF SOME METABIOTICS SYNTHESIZED BY LACTIC ACID BACTERIA

110. Harutyunyan L.

MODERN APPROACHES TO THE SYSTEMIC TREATMENT OF RECURRENT OVARIAN CANCER

119. Absoian T., Hameed Alwan M

ASSOCIATION BETWEEN CAFFEINE, ANXIETY AND THE OCCURRENCE OF APH-THOUS STOMATITIS IN THE ARMENIAN ETHNICITY

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