# Clusters explaining the relation between menopause and self-reported periodontal disease: A cross-sectional study (#107178)

First revision

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# Structure and Criteria



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- 2. EXPERIMENTAL DESIGN
- 3. VALIDITY OF THE FINDINGS
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Conclusions are well stated, linked to original research question & limited to supporting results.

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Τ	p

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Comment on strengths (as well as weaknesses) of the manuscript

## **Example**

Smith et al (J of Methodology, 2005, V3, pp 123) have shown that the analysis you use in Lines 241-250 is not the most appropriate for this situation. Please explain why you used this method.

Your introduction needs more detail. I suggest that you improve the description at lines 57-86 to provide more justification for your study (specifically, you should expand upon the knowledge gap being filled).

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- 1. Your most important issue
- 2. The next most important item
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- 4. The least important points

I thank you for providing the raw data, however your supplemental files need more descriptive metadata identifiers to be useful to future readers. Although your results are compelling, the data analysis should be improved in the following ways: AA, BB, CC

I commend the authors for their extensive data set, compiled over many years of detailed fieldwork. In addition, the manuscript is clearly written in professional, unambiguous language. If there is a weakness, it is in the statistical analysis (as I have noted above) which should be improved upon before Acceptance.



# Clusters explaining the relation between menopause and selfreported periodontal disease: A cross-sectional study

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**Background**. Menopause is an important milestone in the women's life continuum and is associated with potentially adverse effects, including those related to oral health. This study assessed self-reported periodontal disease in relation to menopausal status. **Methods**. A cross-sectional study involving a convenience sample of female university dental hospital attendees was conducted using a validated, self-administered, selfreported periodontal disease questionnaire. A two-step cluster analysis was used to categorize the participants based on menstrual period (MP) continuity, systemic diseases and age. Differences between clusters were analyzed using chi-square test. **Results**. From 112 included participants, 3 clusters resulted from the analysis: Cluster #1 (37±8 years, no systemic diseases and continued MP), Cluster #2 (40±10, with systemic diseases and continued MP) and Cluster #3 (54±9, with systemic diseases and discontinued MP). Cluster #3 had less optimal oral hygiene habits and more missing teeth (p>0.05). Clusters #1 and #2 reported more gingival bleeding, tooth sensitivity and calculus (p>0.05). Cluster #3, on the other hand, presented with more self-reported oral dryness ( $p \le 0.05$ ). **Conclusions**. Within study limits, clusters of menopausal women with systemic diseases reported high symptoms of periodontal disease that were not significantly different from younger individuals, with the exception of oral dryness.

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# **PeerJ**

<del>1</del> 1	Abstract
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#### Introduction

Menopause is defined by the World Health Organization (WHO) as "the permanent cessation of

83 menstruation due to loss of ovarian follicular activity" (World Health Organization, 1996). It is

84 diagnosed retrospectively following 12 months of amenorrhea that is not associated with

85 pathology. The term "Menopause" itself means "without estrogen", and refers to the time at

86 which cyclic ovarian function - known as menstruation - ceases (World Health Organization,

1996). As levels of circulating sex hormones change, several clinical outcomes ensue with a

resulting potential impact on emotional state and quality of life.

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Within the context of oral health, it has been shown that hormonal fluctuations during the different stages of menopause may lead to certain inflammatory responses in the body (Mariotti, 2005). Notably, these fluctuations and changes increase the sensitivity of the gingiva to microbial dental plague and calculus, consequently increasing the susceptibility to adverse oral health outcomes such as gingivitis, periodontitis, oral dysesthesia and xerostomia or dry mouth (Mosley, Smith & Dezan, 2015; "Guideline on Oral Health Care for the Pregnant Adolescent... 2016). Further, the gingival epithelium becomes thinner, atrophic and more prone to inflammatory changes during menopause (Alves et al., 2015). Hormones have been shown to directly and indirectly exert effects on the periodontium. Estrogen and progesterone affect cellular proliferation, differentiation and growth in target tissues, including keratinocytes and fibroblasts in the gingiva (Mariotti & Mawhinney, 2013; Jafri et al., 2015). Therefore, the sudden decrease in estrogen production may be associated with primary osteoporosis, which has an effect on the jaw bones (Alves et al., 2015). The reduction in bone mineral density resulting from osteoporosis contributes to periodontal disease progression in menopausal women (Alves et al., 2015). Moreover, Estrogens may interfere with other periodontal tissues such as the gingiva and periodontal ligament, and influence host immune-inflammatory responses (Alves et al., 2015). Some studies have reported a link between osteoporosis of the mandible and the peripheral skeleton with alveolar bone loss of the mandible and tooth loss in menopausal women (Jonasson & Rythén, 2016).

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Interestingly, periodontal disease is less prevalent in women than men. However, studies have shown that the opposite takes place in older populations (Gupta et al., 2018). The currently increasing life span of populations indicates that the proportion of menopausal women affected by periodontal disease is on the rise. Approximately one in two perimenopausal women is affected by moderate to severe periodontitis, highlighting the need for manual intervention to reduce periodontitis risk (Gil-Montoya et al., 2021).

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Despite the available evidence, there are no data that have focused on the personal input of women in Northwest Saudia Arabia with regards to their oral health, namely periodontal disease and the possible relation with their menopausal condition. This can help oral health promotion, early treatment and prevention of further complications associated with different stages of





121 menopause. Specifically, this will aid in the establishment of a referral system between gynecologists, general practitioners and dentists that involves regular dental checkups for early 122 detection and intervention, and based on the fact that oral health is an integral component of the 123 overall health. 124 125 126 The objective of this study was thus to assess the self-reported periodontal disease among a selected group of women attending a university outpatient dental clinic in relation to their 127 128 menopausal status. 129 **Materials & Methods** 130 131 **Study Design** 132 This was an observational cross-sectional analytical study, conducted at the Taibah University College of Dentistry outpatient clinics in AlMadinah AlMunawwarah, Saudi Arabia. Approval 133 134 from the Taibah University College of Dentistry Research Ethics Committee was obtained prior to commencement of the study, as all set procedural institutional guidelines were followed 135 136 (Approval no. TUCDREC/20180107/Fadl). 137 138 **Ethical Considerations** The study was conducted in accordance with the ethical guidelines set by the Declaration of 139 Helsinki (2013). Before recruitment, the purpose of the study was explained to each of the 140 141 participants and informed consent was obtained. They were also informed that participation was 142 voluntary, with no negative repercussions in terms of quality of offered healthcare due to declining to participate. Participants were also assured that all information will be kept 143 confidential and will only be used for research and educational purposes. In addition, participants 144 145 were informed whenever necessary of their treatment needs based on the findings of this study. 146 147 **Study Population and Sample Size Calculation** The study involved a convenience / consecutive sample from all female individuals aged 148 between 22 and 75 years, who were attending the college clinics between November 2018 and 149 150 July 2019. Children and individuals below 22 years were excluded alongside those with 151 dementia or mental health or radiotherapy. Approximately 13,000 patients (women and men) 152 visit the clinics each year.

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157 158 Based on the reported prevalence of the outcome "gingivitis" in the general population of 75%, with a desired level of confidence of 95%, and a margin of error of 10, and an estimated population of women attending the clinics annually of 6,500, a sample size of 97 participants was required for the study (Chisquares Inc. Sample Size Calculation for Cross-sectional Studies with Percentage as Outcome.). Incomplete responses were to be excluded from the analysis.



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#### **Interview and Questionnaire**

- Data were collected by means of an interview and a self-administered questionnaire. The
- interview was conducted in the waiting area and involved explanation of the study, giving
- 164 consent to participate and filling the questionnaire. The questionnaire took 3-5 minutes to
- 165 complete and included providing information about age, education level, marital status, siblings,
- current health status and social and oral hygiene habits. Participants were also asked about
- 167 continuation of their menstrual period (MP). Moreover, a valid and reliable Arabic version of the
- self-reported periodontal health questionnaire was included (Khader, Alhabashneh & Alhersh,
- 169 2014). This section consisted of seventeen close ended dichotomous questions, representing the
- 170 outcome variables.

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### **Data Analysis**

- 173 A statistical package (SPSS Inc., Chicago, IL, USA version 20) was used to calculate
- 174 descriptive values in the form of means and standard deviations for continuous variables, and
- 175 frequencies/percentages for categorical variables. A 2-step cluster analysis was performed to
- identify hidden patterns and relationships by categorizing participants based on the reported
- background, demographic and health data. Repeated attempts utilizing different demographic
- and habitual variables produced a clustering model of "Good" quality via Silhouette's measure.
- 179 The finally selected clustering predictors, in order of importance; were menstrual period
- 180 continuation, presence of systemic diseases and age. A post hoc power calculation was
- performed considering the clusters with the highest (n=65) and lowest (n=22) number of
- participants and a confidence interval of 95% yielded a study power of 77% (OpenEpi, Version
- 183 3, open source calculator PowerCross). The differences between groups/clusters in terms of
- quantitative demographics, oral hygiene habits, oral findings and the self-reported periodontal
- disease questions were analyzed using the chi-square test. A p-Value  $\leq 0.05$  was considered
- 186 statistically significant.

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#### Results

- 189 Of a total of 114 participating female patients, two were excluded due to incomplete data. The
- remaining 112 returned their completed questionnaires and agreed to participate in the study
- 191 while awaiting their treatment appointment at the dental clinics. The mean age of the total
- sample was  $40 (\pm 10)$  years. Fifty-two percent held a university degree and 66% were employed.
- 193 Thirty-six percent of the participants suffered from different systemic conditions such as
- hypertension and diabetes mellitus. Similarly, 59% were on non-specified chronic medication,
- 195 five of which specifically reporting the use of hormone replacement therapy (HRT). Fourteen
- 196 percent of the total participants were current smokers, 3 of them smoked cigarettes and the
- 197 remaining used waterpipes. Twenty percent had already discontinued their menstrual period.

- The two-step cluster analysis resulted in 3 clusters, with cluster # 1 (n=65) including women
- 200 with no systemic conditions and cluster # 3 (n=22) involving those with a completely



discontinued menstrual period (Figure 1). Compared to the other two clusters, women in cluster # 3 mostly completed up to high school education (p < 0.001), 55% were unemployed (p < 0.05) and had the highest number of children (p < 0.01) (Table 1). With regards to participants from cluster # 2, on the other hand, 84% were married (p > 0.05) and 80% were on medication (p < 0.01) (Table 1).

Cluster # 3 apparently had the least percentage of participants who regularly brushed their teeth (91%), used other tooth cleaning aids (41%) or used mouthwash (41%) compared to the other two clusters, although the differences were not statistically significant (p > 0.05) (Figure 2). The same cluster also tended to have the highest prevalence of tooth loss (96%) and replacement of missing teeth (50%) compared to the other clusters (Figure 3). However, the differences were insignificant (p > 0.05). With regards to the self-reported periodontal disease (SRPD) questions, the observations of swollen and painful gums, pus, movable teeth, teeth that have changed in position or longer teeth were more reported among individuals in cluster # 3, albeit the lack of statistical significance (p > 0.05) (Table 2). The feeling of dry mouth was also significantly higher in cluster # 3 compared to the other 2 clusters (p < 0.05) (Table 2). On the other hand, self-reported bleeding gums, tooth sensitivity and calculus deposits were relatively, yet

insignificantly; higher in clusters # 1 and # 2 (p > 0.05) (Table 2).

## **Discussion**

This study aimed at evaluating self-reported oral health in women in relation to their menopausal status. Women who have not reached menopausal age insignificantly reported more bleeding gums, tooth sensitivity and calculus deposits than their menopausal counterparts. This is in line with findings from larger registries, reporting more oral health problems by younger women (Azofeifa et al., 2014). An explanation could be that younger women are exposed to continuous hormonal changes, with the potential of exaggerated responses of the dental and periodontal tissues to local stimuli (Dar-Odeh et al., 2017). Furthermore, the fact that women in the reproductive age may go through several pregnancies, are minimally aware at the time with regards to their oral health and often avoid dental visits could all contribute to the characteristic adverse oral health findings (Dar-Odeh et al., 2018).

There was an insignificant tendency for older menopausal women with different systemic comorbidities, i.e. Cluster # 3, to observe more painful gums, pus, longer teeth, teeth that are movable, or teeth that have changed their position, all of which are known symptoms related to advanced periodontitis. Recent cohort studies showed that postmenopausal women were more likely to have periodontitis (Park et al., 2023a), and that women with periodontitis are more likely to develop osteoporosis (Choi et al., 2017). Furthermore, it was concluded from previous studies that diabetes mellitus and hypertension are particularly associated with missing teeth among women (Dar-Odeh et al., 2019). A recent nested case-control study showed that healthy patients with periodontitis had a higher mean systolic and diastolic blood pressure than





periodontitis-free individuals (Muñoz Aguilera et al., 2021). The increased local and systemic inflammatory markers detected in periodontitis patients are associated with vascular changes and endothelial dysfunction. Furthermore, several studies concluded that diabetic patients had a higher prevalence of periodontal disease than healthy individuals, and attributed the development of diabetes to oral infection and periodontal pathogens (Martínez-García & Hernández-Lemus, 2021).

It was important to investigate the potential role of smoking in this study. Smoking is considered a major risk factor in periodontal disease development and progression. Recent research suggests that smoking acts as a facilitator for the colonization of periodontal pathogens, influencing the structure of the subgingival microbial community and worsening treatment outcomes (Jiang et al., 2020). However, no differences in smoking was observed between the three studied clusters.

Additional findings from this study confirmed what was mentioned earlier, which showed a tendency for menopausal women to have higher prevalence of missing teeth and tooth replacement. Interestingly, the current study identified relatively poor oral hygiene practices among menopausal women, which aggravates the risk for dental caries and periodontal disease. Moreover, menopausal women were of low educational backgrounds, mostly unemployed and with more children. These sociodemographic variables could play a role in predisposing to low awareness to oral hygiene practices and favorable oral health outcomes. Park and co-workers concluded in their nationwide cohort study that good oral hygiene practices are adversely correlated with osteoporotic fractures (Park et al., 2023b).

Menopausal women with systemic comorbidities reported a significantly higher prevalence of dry mouth. Krupa et al. have recently concluded that there was a significant association between menopausal duration and salivary flow rates (Krupa et al., 2023). Older age is associated with polypharmacy which could also predispose to dry mouth (Cannon et al., 2023). On the other hand, several medications are known for improving periodontal health. Non-steroidal anti-inflammatory drugs are thought to enhance periodontal treatment outcomes through their anti-inflammatory effects (Ren et al., 2023). Further, hormone replacement therapy (HRT) during menopause is thought to relieve dry mouth symptoms (Wang et al., 2021). Moreover, HRT seems to enhance alveolar bone density and teeth durability (Pizzo et al., 2011). However, as low as only five participants reported the use of HRT in the current study, which statistically limits its influence on the study outcomes.

The relatively small sample size is looked upon as a study limitations, since it may have impacted the extrapolation of the observed findings alongside the convenience sampling that precluded generalizability of the finding. This may dictate the consideration of the current investigation as a pilot study at best. However, the self-reporting of oral health symptoms in menopausal women in this study covers an important aspect, providing unique information that





can be utilized for hypothesis generation in future studies. Moreover, the cross-sectional design limits the identification of temporal relationships between menopause and oral health parameters, necessitating the interpretation of findings with caution. Another limitation is the non-disclosure of the medication types consumed by more than half of the participants, which in theory may have included drugs that increase (anti-cancer drugs) or decrease (anti-inflammatory agents) the risk of periodontitis, although minimal conclusive evidence on such associations is available in the literature (Albandar, Susin & Hughes, 2018).

### **Conclusions**

From this investigation, it can be concluded that clusters of menopausal women with systemic diseases reported high symptoms of oral and periodontal disease that were not significantly different from younger individuals, with the exception of oral dryness. Further studies with larger study samples are required to expand on such findings.



#### 318 **References**

- 319 Albandar JM, Susin C, Hughes FJ. 2018. Manifestations of systemic diseases and conditions that
- affect the periodontal attachment apparatus: Case definitions and diagnostic considerations.
- *Journal of clinical periodontology* 45 Suppl 20:S171–S189. DOI: 10.1111/jcpe.12947.
- 322 Alves RC, Félix SA, Rodriguez-Archilla A, Oliveira P, Brito J, Dos Santos JM. 2015.
- Relationship between menopause and periodontal disease: a cross-sectional study in a
- Portuguese population. *International journal of clinical and experimental medicine* 8:11412–
- 325 11419.
- 326 Azofeifa A, Yeung LF, Alverson CJ, Beltrán-Aguilar E. 2014. Oral health conditions and dental
- visits among pregnant and nonpregnant women of childbearing age in the United States,
- National Health and Nutrition Examination Survey, 1999-2004. Preventing chronic disease
- 329 11:E163. DOI: 10.5888/pcd11.140212.
- Cannon I, Robinson-Barella A, McLellan G, Ramsay SE. 2023. From Drugs to Dry Mouth: A
- 331 Systematic Review Exploring Oral and Psychological Health Conditions Associated with Dry
- Mouth in Older Adults with Polypharmacy. Drugs & aging 40:307–316. DOI: 10.1007/s40266-
- 333 023-01017-5.
- Choi J-K, Kim Y-T, Kweon H-I, Park E-C, Choi S-H, Lee J-H. 2017. Effect of periodontitis on
- the development of osteoporosis: results from a nationwide population-based cohort study
- 336 (2003-2013). BMC women's health 17:77. DOI: 10.1186/s12905-017-0440-9.
- 337 Dar-Odeh NS, Aleithan FA, Alnazzawi AA, Al-Shayyab MH, Abu-Hammad SO, Abu-Hammad
- OA. 2017. Factors affecting oral health determinants in female university students: a cross-
- 339 sectional survey in Saudi Arabia. *International journal of adolescent medicine and health*
- 340 32:/j/ijamh.2020.32.issue-1/ijamh-2017-0084/ijamh-2017-0084.xml. DOI: 10.1515/ijamh-
- 341 2017-0084.
- Dar-Odeh NS, Binsaad S, Gasim R, Haidary R, Afandi A, Hammad S, Hammad O. 2018. Why
- do Women avoid Dental Visits During Pregnancy? A CrossSectional Survey in Al Madinah,
- Western Saudi Arabia. Pesquisa Brasileira em Odontopediatria e Clínica Integrada 18:1–8.
- 345 DOI: 10.4034/pboci.2018.181.31.
- 346 Dar-Odeh NS, Borzangy S, Babkair H, Farghal L, Shahin G, Fadhlalmawla S, Alhazmi W, Taher
- S, Abu-Hammad O. 2019. Association of Dental Caries, Retained Roots, and Missing Teeth
- with Physical Status, Diabetes Mellitus and Hypertension in Women of the Reproductive Age.
- 349 *International journal of environmental research and public health* 16. DOI:
- 350 10.3390/ijerph16142565.
- 351 Gil-Montoya JA, Garrido-Martínez M, Barrios-Rodríguez R, Ramos-García P, Lenouvel D,
- 352 Montes-Castillo C, Martínez-Ramírez MJ. 2021. Association between low bone mineral
- density and periodontitis in generally healthy perimenopausal women. *Journal of*
- *periodontology* 92:95–103. DOI: 10.1002/JPER.20-0029.
- 355 Guideline on Oral Health Care for the Pregnant Adolescent. 2016. *Pediatric dentistry* 38:59–66.
- 356 Gupta N, Sprouse L, Sadeghi G, Thacker H, Williams K, Palomo L. 2018. Social Determinants
- of Health and Periodontitis in Postmenopausal Women. *J Dents Dent Med* 1:106.



- 358 Jafri Z, Bhardwaj A, Sawai M, Sultan N. 2015. Influence of female sex hormones on
- periodontium: A case series. *Journal of natural science, biology, and medicine* 6:S146-149.
- 360 DOI: 10.4103/0976-9668.166124.
- 361 Jiang Y, Zhou X, Cheng L, Li M. 2020. The Impact of Smoking on Subgingival Microflora:
- From Periodontal Health to Disease. Frontiers in microbiology 11:66. DOI:
- 363 10.3389/fmicb.2020.00066.
- Jonasson G, Rythén M. 2016. Alveolar bone loss in osteoporosis: a loaded and cellular affair?
- *Clinical, cosmetic and investigational dentistry* 8:95–103. DOI: 10.2147/CCIDE.S92774.
- 366 Khader Y, Alhabashneh R, Alhersh F. 2014. Validity of a Self-Reported Periodontal Disease
- 367 Questionnaire among Jordanians. 4:2–7. DOI: 10.4172/2161-1122.100020.
- 368 Krupa NC, Thippeswamy HM, Chandrashekar BR, Thetakala RK. 2023. Impact of Menopausal
- Duration on Salivary Flow Rate, Tooth Loss, and Oral Health-related Quality of Life in Indian
- 370 Communities. Journal of mid-life health 14:101–106. DOI: 10.4103/jmh.jmh 41 22.
- 371 Mariotti AJ. 2005. Estrogen and extracellular matrix influence human gingival fibroblast
- proliferation and protein production. *Journal of periodontology* 76:1391–1397. DOI:
- 373 10.1902/jop.2005.76.8.1391.
- Mariotti A, Mawhinney M. 2013. Endocrinology of sex steroid hormones and cell dynamics in
- 375 the periodontium. *Periodontology 2000* 61:69–88. DOI: 10.1111/j.1600-0757.2011.00424.x.
- 376 Martínez-García M, Hernández-Lemus E. 2021. Periodontal Inflammation and Systemic
- 377 Diseases: An Overview. *Frontiers in physiology* 12:709438. DOI: 10.3389/fphys.2021.709438.
- 378 Mosley JF 2nd, Smith LL, Dezan MD. 2015. An overview of upcoming changes in pregnancy
- and lactation labeling information. *Pharmacy practice* 13:605. DOI:
- 380 10.18549/pharmpract.2015.02.605.
- 381 Muñoz Aguilera E, Suvan J, Orlandi M, Miró Catalina Q, Nart J, D'Aiuto F. 2021. Association
- 382 Between Periodontitis and Blood Pressure Highlighted in Systemically Healthy Individuals:
- Results From a Nested Case-Control Study. *Hypertension (Dallas, Tex. : 1979)* 77:1765–1774.
- 384 DOI: 10.1161/HYPERTENSIONAHA.120.16790.
- Park K-Y, Kim M-H, Choi S-H, Pang E-K. 2023a. Association of periodontitis with menopause
- and hormone replacement therapy: a hospital cohort study using a common data model. *Journal*
- *of periodontal & implant science* 53:184–193. DOI: 10.5051/jpis.2202480124.
- Park J-H, Park M-S, Kim H-J, Lee H, Kim J-W, Song T-J. 2023b. Better oral hygiene is
- associated with a reduced risk of osteoporotic fracture: a nationwide cohort study. Frontiers in
- 390 *endocrinology* 14:1253903. DOI: 10.3389/fendo.2023.1253903.
- 391 Pizzo G, Guiglia R, Licata ME, Pizzo I, Davis JM, Giuliana G. 2011. Effect of hormone
- replacement therapy (HRT) on periodontal status of postmenopausal women. *Medical science*
- 393 *monitor*: international medical journal of experimental and clinical research 17:PH23-27.
- 394 DOI: 10.12659/msm.881700.
- Ren J, Fok MR, Zhang Y, Han B, Lin Y. 2023. The role of non-steroidal anti-inflammatory
- drugs as adjuncts to periodontal treatment and in periodontal regeneration. *Journal of*
- 397 translational medicine 21:149. DOI: 10.1186/s12967-023-03990-2.



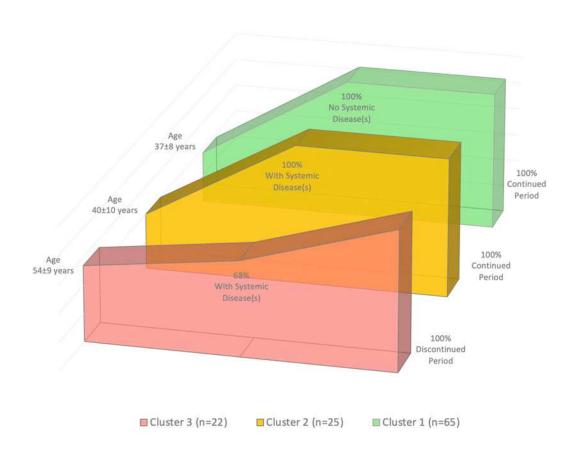
# **PeerJ**

398	Wang L, Zhu L, Yao Y, Ren Y, Zhang H. 2021. Role of hormone replacement therapy in
399	relieving oral dryness symptoms in postmenopausal women: a case control study. BMC oral
400	health 21:615. DOI: 10.1186/s12903-021-01966-6.
401	World Health Organization. 1996. Research on the menopause in the 1990s: report of a WHO
402	scientific group. Geneva, Switzerland: WHO.
403	World Medical Association Declaration of Helsinki: Ethical Principles for Medical Research
404	Involving Human Subjects. 2013. JAMA 310:2191–2194.
405	



# Figure 1

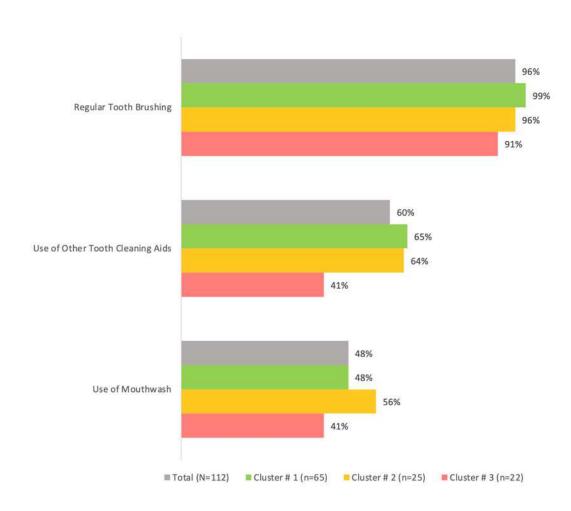
An area chart showing the three resulting clusters from the 2-step cluster analysis, with the predictors "Period Continuation", "Presence of Systemic Diseases" and "Age".





# Figure 2

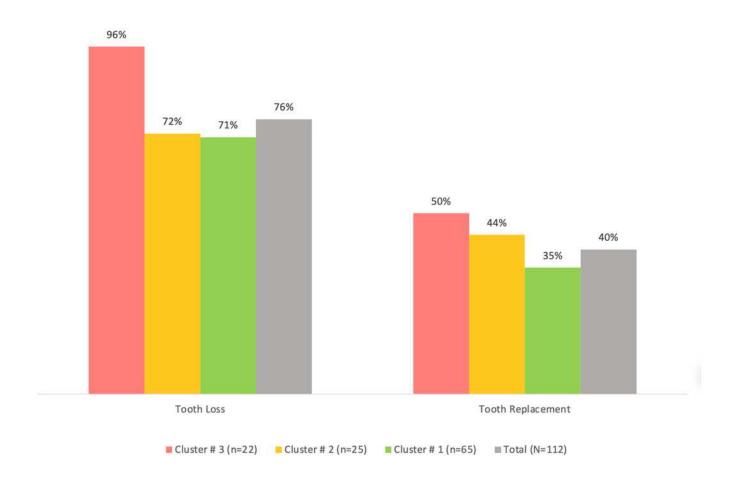
A bar-chart showing the oral hygiene habits in the total sample (N=112) and the three clusters. No significant differences were observed using chi-square test.





# Figure 3

A bar-chart showing the prevalence of self-reported tooth loss and tooth replacement in the total sample (N=112) and the three clusters. No significant differences were observed using chi-square test.





## Table 1(on next page)

Demographic characteristics of the total sample (N=112) and the three clusters.

(p-Values in **BOLD** fonts are statistically significant using chi-square test at 0.05). <sup>a</sup> Only 5 (5%) of those on medication are on hormone replacement therapy (HRT). <sup>b</sup> Only 3 (3%) smoke cigarettes, the remaining smokers use water pipe.

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Table 1: Demographic characteristics of the total sample (N=112) and the three clusters.

Variable	Total Sample (N=112)	Cluster # 1 (n=65)	Cluster # 2 (n=25)	Cluster # 3 (n=22)	p Value
Marital Status - n (%)					
Not Married	27 (24)	18 (28)	4 (16)	5 (23)	0.502
Married	85 (76)	47 (72)	21 (84)	17 (77)	
Education - n (%)					
High school or lower	54 (48)	26 (40)	12 (48)	20 (91)	0.000
University degree	58 (52)	39 (60)	13 (52)	2 (9)	
Employment - n (%)					
No	38 (34)	22 (34)	4 (16)	12 (55)	0.021
Yes	74 (66)	43 (66)	21 (84)	10 (45)	
Siblings - n (%)					
None	27 (24)	17 (26)	4 (16)	6 (27)	
1-2	22 (20)	17 (26)	4 (16)	1 (5)	0.001
3-4	33 (29)	19 (29)	12 (48)	2 (9)	
> 4	30 (27)	12 (19)	5 (20)	13 (59)	
Recent Medication - n (%) *					
No	46 (41)	35 (54)	5 (20)	6 (27)	0.005
Yes	66 (59)	30 (46)	20 (80)	16 (73)	
Smoking - n (%) †					
No	96 (86)	53 (82)	22 (88)	21 (96)	0.255
Yes	16 (14)	12 (18)	3 (12)	1 (4)	

(p-Values in **BOLD** fonts are statistically significant using chi-square test at 0.05)

<sup>\*</sup> Only 5 (5%) of those on medication are on HRT

<sup>†</sup> Only 3 (3%) smoke cigarettes, the remaining smokers use water pipe



# Table 2(on next page)

Positive answers (YES) to the self-reported periodontal disease questionnaire by the total sample (N=112) and the three clusters.

(p-Values in **BOLD** fonts are statistically significant using chi-square test at 0.05)



Table 2: Positive answers (YES) to the self-reported periodontal disease questionnaire by the total sample (N=112) and the three clusters.

	Total	Cluster #	Cluster #	Cluster #	n	
Variable	Sample	1	2	3	p Value	
	(N=112)	(n=65)	(n=25)	(n=22)	value	
Did you notice any swollen or reddish area in						
your gums (more than the usual)?	30 (27)	17 (26)	6 (24)	7 (32)	0.820	
Do your gums bleed easily?	50 (45)	30 (46)	12 (48)	8 (36)	0.676	
Do you feel pain from your gums?	28 (25)	12 (18)	8 (32)	8 (36)	0.161	
Does your mouth feel dry?	29 (26)	11 (17)	8 (32)	10 (45)	0.022	
Have you noticed an unpleasant smell from						
your mouth?	35 (31)	24 (37)	5 (20)	6 (27)	0.271	
Do you have any abscess or pus in your mouth?	13 (12)	7 (11)	1 (4)	5 (23)	0.128	
Does food get trapped between your teeth?	85 (76)	51 (79)	18 (72)	16 (73)	0.755	
Do you feel that your teeth move / have						
moved?	15 (13)	5 (8)	4 (16)	6 (27)	0.060	
Have you noticed that your teeth have changed						
in their position?	23 (20)	12 (18)	5 (20)	6 (27)	0.674	
Have you noticed that your teeth have become						
longer?	18 (16)	8 (12)	4 (16)	6 (27)	0.255	
Do you feel any sensitivity in your teeth?	46 (41)	28 (43)	13 (52)	5 (23)	0.111	
Do you have any calculus or limestone deposits						
on your teeth?	60 (54)	33 (51)	16 (64)	11 (50)	0.494	
Do you think you have any disease in your						
gums or tissues that support your teeth						
(periodontium) or loss of bone that fixes the						
teeth?	18 (16)	9 (14)	4 (16)	5 (23)	0.618	
Has any dentist or hygienist ever told you that						
you have deep periodontal pockets?	6 (5)	3 (5)	2 (8)	1 (4)	0.801	
Have you ever been told that you needed						
treatment for your gums / periodontal tissues?	16 (14)	7 (11)	3 (12)	6 (27)	0.150	
Have you ever received any treatment for your						
gums / periodontal tissues?	26 (23)	13 (20)	6 (24)	7 (32)	0.522	
Have you ever received any surgical treatment						
for your gums / periodontal tissues?	3 (3)	3 (5)	0 (0)	0 (0)	0.328	
dues in <b>ROLD</b> fonts are statistically significant using chi-square test at 0.05)						

(p-Values in **BOLD** fonts are statistically significant using chi-square test at 0.05)



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