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Retrospective Study

Establishment of models to predict factors influencing periodontitis in patients with type 2 diabetes mellitus

Xu HM *et al.* Study of periodontitis in T2DM

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Abstract

BACKGROUND

Type 2 diabetes mellitus (T2DM) is associated with periodontitis. Currently, there are few studies proposing predictive models for periodontitis in patients with T2DM.

AIM

To determine the factors influencing periodontitis in patients with T2DM by constructing logistic regression and random forest models.

METHODS

In this a retrospective study, 300 patients with T2DM who were hospitalized at the First People's Hospital of Wenling from January 2022 to June 2022 were selected for inclusion, and their data were collected from hospital records. We used logistic regression to analyze factors associated with periodontitis in patients with T2DM, and random forest and logistic regression prediction models were established. The prediction efficiency of the models was compared using the area under the receiver operating characteristic curve (AUC).

RESULTS

Of 300 patients with T2DM, 224 had periodontitis, with an incidence of 74.67%. Logistic regression analysis showed that age [odds ratio (OR) = 1.047, 95% confidence interval (CI): 1.017-1.078], teeth brushing frequency (OR = 4.303, 95%CI: 2.154-8.599), education level (OR = 0.528, 95%CI: 0.348-0.800), glycosylated hemoglobin (HbA1c) (OR = 2.545, 95%CI: 1.770-3.661), total cholesterol (TC) (OR = 2.872, 95%CI: 1.725-4.781), and triglyceride (TG) (OR = 3.306, 95%CI: 1.019-10.723) influenced the occurrence of periodontitis ($P < 0.05$). The random forest model showed that the most influential variable was HbA1c followed by age, TC, TG, education level, brushing frequency, and sex. Comparison of the prediction effects of the two models showed that in the training dataset, the AUC of the random forest model was higher than that of the logistic

regression model (AUC = 1.000 vs AUC = 0.851; $P < 0.05$). In the validation dataset, there was no significant difference in AUC between the random forest and logistic regression models (AUC = 0.946 vs AUC = 0.915; $P > 0.05$).

CONCLUSION

Both random forest and logistic regression models have good predictive value and can accurately predict the risk of periodontitis in patients with T2DM.

Key Words: Type 2 diabetes mellitus; Periodontitis; Logistic regression; Prediction model; Random forest model; Gingival disease

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Core Tip: With the rapid increase in the number of patients with type 2 diabetes mellitus (T2DM), the number of cases complicated by periodontitis has also increased. Without timely intervention, periodontitis can lead to tooth loosening and loss, and a decline in oral function, reducing patient quality of life. We retrospectively analyzed the data of 300 patients with T2DM to determine the factors influencing periodontitis. Random forest and logistic regression models were constructed to provide a theoretical basis for predicting periodontitis in patients with T2DM.

INTRODUCTION

Type 2 diabetes mellitus (T2DM) accounts for more than 90% of cases of diabetes, and occurs mostly in adults over 40 years of age^[1,2]. Research has shown that individuals with diabetes are more likely to develop periodontitis than those without^[3]. Periodontitis is a chronic inflammation that occurs in periodontal supporting tissues and is characterized by gingival inflammation, formation of a periodontal pocket, resorption and destruction of the alveolar bone, and tooth loosening, displacement, and loss^[4]. It may also affect

masticatory function and nutritional intake^[5]. Many studies have shown a bidirectional relationship between T2DM and periodontitis, and the incidence of periodontitis in patients with T2DM is approximately 2-3 times that of the general population^[6]. Many studies have reported on the factors influencing periodontitis in patients with T2DM, but the majority use logistic regression analysis, which cannot intuitively present the importance of the outcome^[7,8]. With the advent of the era of big data, predictive models have become useful in predicting the occurrence of diseases, but there are few relevant prediction models for periodontitis in patients with T2DM. Studies have shown that among the multiple machine learning models for predicting the risk of kidney disease in patients with T2DM, the random forest model has better performance and higher accuracy than logistic regression^[9]. Therefore, the objective of this study was to retrospectively analyze the factors influencing periodontitis in patients with T2DM, and establish random forest and logistic regression prediction models for this disease.

MATERIALS AND METHODS

Subjects

Three hundred patients with T2DM who were hospitalized at the First People's Hospital of Wenling from January 2022 to June 2022 were selected as research subjects, and their relevant data were collected for this retrospective study (Figure 1). The inclusion criteria were: (1) Age ≥ 18 years old; and (2) T2DM diagnosed at our hospital and without other serious complications. The exclusion criteria were: (1) Patients with other systemic diseases affecting periodontal health; (2) Serious organ disorders; (3) Cognitive or mental disorders; (4) Periodontal treatment in the past 3 mo; (5) Pregnancy; (6) Unable to accept oral periodontal examination; and (7) Incomplete data.

Methods

General information and clinical examination data were collected from hospital records. General patient information included monthly income, age, education level, body mass index (BMI), sex, smoking, and alcohol intake. We also asked about daily brushing

frequency and exercise habits. Clinical examination data included glycosylated hemoglobin (HbA1c), total cholesterol (TC), triglyceride (TG), high-density lipoprotein cholesterol (HDL-C), and low-density lipoprotein cholesterol (LDL-C) levels.

Diagnostic criteria and classification of periodontitis

The Diagnostic criteria for periodontitis were: (1) A definite history of periodontitis; (2) Loose teeth in the mouth due to periodontitis without other factors; and (3) Periodontal examination showing that the depth of a periodontal pocket of at least one index tooth was ≥ 4 mm, and the tooth had clinical attachment loss (AL) ≥ 1 mm. According to the diagnostic criteria for periodontitis, the patients were divided into periodontitis and no periodontitis groups. Based on the depth of the periodontal pocket, AL, and alveolar bone resorption, periodontitis was divided into three categories: (1) Mild: Gingival inflammation and bleeding on probing, probing depth of periodontal pocket ≤ 4 mm, AL 1-2 mm, and alveolar bone resorption not more than one third of root length; (2) Moderate: Gingival inflammation, bleeding on probing or pus, depth of periodontal pocket ≤ 6 mm, AL 3-4 mm, alveolar bone resorption more than one third of the length of the root, but not more than half the length of the root, mild tooth loosening, and mild furcation lesions; and (3) Severe: Obvious inflammation accompanied by periodontal abscess, periodontal pocket depth > 6 mm, AL ≥ 5 mm, alveolar bone resorption more than half of the root length, multiple root furcation lesions, and tooth loosening and displacement.

Classification of indicators

According to BMI [weight (kg)/height (m²)], the patients were classified as: Underweight: BMI < 18.5 ; normal weight: BMI 18.5-24.0; overweight: BMI 24.0-28; obesity: BMI ≥ 28 . According to HbA1c, blood glucose control was classified as: Ideal control: HbA1c $< 6.5\%$; good control: HbA1c 6.5%-7.5%; poor control: HbA1c 7.5%-8.5%; very poor control: HbA1c $\geq 8.5\%$. The normal levels of TC, TG, HDL-C, and LDL-C are < 5.18 mmol/L, < 1.7 mmol/L, ≥ 1.04 mmol/L, and ≤ 3.37 mmol/L, respectively.

Statistical analysis

Patient information was analyzed using SPSS 26.0. For univariate analysis, the data that passed the normality test, represented by the mean \pm SD, were compared by the *t*-test. Data that failed the normality test, presented as median (M) and 25% and 75% percentiles (P25, P75), were compared by the rank sum test. Count data, denoted as *n* (%), were compared by the χ^2 test. Statistical significance was set at $P < 0.05$. The variables that were statistically significant in the univariate analysis were included in the logistic regression to analyze the factors influencing periodontitis in T2DM patients. Logistic regression and random forest prediction models were constructed using the R, and the receiver operating characteristic curves of the training dataset and validation dataset of the two models were drawn. The predictive efficacy of the models was compared according to the area under curve (AUC), and the AUC was compared using the Delong test.

RESULTS

Analysis of general information and periodontitis

Among the 300 patients with T2DM, 224 had periodontitis, with an incidence of 74.67%. Among them, 83 (37.05%) had mild, 78 (34.82%) had moderate, and 63 (28.13%) had severe periodontitis. One hundred and sixty-four (54.67%) were male and 136 (45.33%) were female, with an incidence based on sex of 58.04% and 44.74%, respectively. The average age was 60.91 ± 9.49 years in the periodontitis group and 54.11 ± 14.69 years in the no periodontitis group. The proportion of patients with education level below high school was highest (43.75%) in the periodontitis group. In terms of monthly income, the highest proportion of monthly income in the periodontitis group was between 2000 and 5000 yuan (44.20%). Univariate analysis showed that there were significant differences in sex, age, and education level between the periodontitis and non-periodontitis groups ($P < 0.05$), but there was no significant difference in monthly income ($P > 0.05$; Table 1).

Analysis of lifestyle and periodontitis

Of the patients with periodontitis, 23.21% smoked, 17.86% drank alcohol, 51.34% exercised regularly, and 63.39% brushed their teeth less than twice a day. Of the patients without periodontitis, 13.16% smoked, 14.47% drank alcohol, 56.58% exercised regularly, and 27.63% brushed their teeth less than twice a day. Univariate analysis showed that there was a significant difference in brushing frequency between the periodontitis and non-periodontitis groups ($P < 0.05$), but there were no significant differences in smoking, drinking, or regular exercise ($P > 0.05$; Table 2).

Analysis of physical health status and periodontitis

Among the 300 patients with T2DM, the overall BMI was 24.96 (22.36, 26.81). The BMI of patients with periodontitis was overweight at 24.86 (22.13, 26.65), and the BMI of patients without periodontitis was overweight at 25.61 (23.26, 27.23). There were 72.77%, 64.29%, and 58.48% of patients in the periodontitis group with normal TG, TC, and HDL-C, respectively. These were lower than the proportions of patients without periodontitis with normal TG, TC, and HDL-C. All patients had normal levels of LDL-C. The periodontitis group had the highest proportion of patients with very poor blood glucose control at 37.50%. The proportion of patients in the non-periodontitis group with good blood glucose control was 40.79%. Univariate analysis showed that there were significant differences in TG, TC, and HbA1c between the two groups ($P < 0.05$), but there were no significant differences in BMI, LDL-C, or HDL-C ($P > 0.05$; Table 3).

Multivariate logistic regression analysis

Whether patients with T2DM developed periodontitis (no occurrence = 0, occurrence = 1) was used as the dependent variable, and statistically significant variables (sex, age, brushing frequency, education level, HbA1c, TC, and TG) were used as independent variables in the univariate analysis (Table 4). Logistic regression analysis showed that age, brushing frequency, education level, HbA1c, TC, and TG were factors influencing periodontitis in patients with T2DM ($P < 0.05$). Older age was associated with a higher risk of periodontal disease [odds ratio (OR) = 1.047, 95% confidence interval (CI): 1.017-

1.078]. Brushing teeth < 2 times/d was also associated with a higher risk of periodontal disease (OR = 4.303, 95%CI: 2.154-8.599). The higher the education level, the lower the risk of periodontitis (OR = 0.528, 95%CI: 0.348-0.800). Higher HbA1c, TC, and TG levels were associated with a higher risk of periodontal disease (OR = 2.545, 95%CI: 1.770-3.661, OR = 2.872, 95%CI: 1.725-4.781, OR = 3.306, 95%CI: 1.019-10.723, respectively; Table 5).

Random forest model

According to the random number method, patients were randomly divided into the training dataset ($n = 200$) and validation dataset ($n = 100$) according to 2/3 and 1/3 of the total number of patients, respectively. The seven variables (age, brushing frequency, HbA1c, TC, education level, TG, and gender) that were statistically significant in the univariate analysis were included in the random forest model. As shown in Figure 2, the importance of variables influencing the occurrence of periodontitis in patients with T2DM was ranked as HbA1c, age, TC, TG, educational level, brushing frequency, and sex.

Comparison between the random forest and logistic regression models

In the training dataset, the overall efficacy of the random forest model in predicting periodontitis in patients with T2DM was higher than that of the logistic regression model. The AUC of the random forest model was significantly higher than that of the logistic regression model ($P < 0.05$; Figure 3A, Table 6). In the validation dataset, the overall performances of the random forest and logistic regression models were comparable and there was no significant difference in AUC between them ($P > 0.05$; Figure 3B, Table 7).

DISCUSSION

Periodontitis is internationally recognized as the sixth most common complication of diabetes. With an increase in the number of diabetic patients, the number of people with periodontitis in the diabetic population is also increasing^[10-12]. The results of this study showed that among 300 patients with T2DM treated at our hospital, 74.67% had periodontitis, which is higher than that reported by de Miguel-Infante *et al*^[13] (23.8%) and

Hong *et al*^[14] (43.7%). The differences in the incidence of periodontitis may be linked to population differences in different countries and regions or to the lifestyle of people in different regions. The higher incidence of periodontitis in patients with T2DM in this study may be due to the older age of the study population as a whole and the irreversible damage to the periodontal tissue caused by inflammation^[15].

Our study found that periodontitis in patients with T2DM was associated with several factors. Logistic regression analysis showed that age, brushing frequency, education level, HbA1c, TC, and TG were the factors significantly influencing periodontitis in patients with T2DM. The importance of the variables was ranked by the random forest model as HbA1c, age, TC, TG, education level, brushing frequency, and sex. The results of the two models were similar, indicating that the prediction results were reliable.

We found that the greater the HbA1c level, the higher the risk of periodontitis (OR = 2.545, 95%CI: 1.770-3.661), which is similar to the results of Wu *et al*^[16], Qureshi *et al*^[17], and Dhir *et al*^[18]. HbA1c is reflective of a patient's blood glucose control in the past 2-3 mo. Poor control of blood glucose can cause more severe periodontitis symptoms, while good blood glucose control can delay the progression of periodontitis^[19,20]. We also found that the risk of periodontal disease increased with age (OR = 1.047, 95%CI: 1.017-1.078). This is similar to the results of a study by de Miguel-Infante *et al*^[13]. Age may be associated with disease as with increasing age, there is generally a weakened immune system making tissues more susceptible to the invasion of anaerobic bacteria and aggravating susceptibility to periodontitis^[21]. Blood lipid levels, particularly higher levels of TC and TG, were associated with an increased risk of periodontal disease. This is similar to the results obtained by Dhir *et al*^[18] and Ding *et al*^[22]. High blood lipid levels in patients with T2DM may increase the body's susceptibility to periodontitis by promoting the expression of inflammatory factors, increasing the oxidative stress response and lipid peroxidation, inhibiting bone formation, and promoting bone resorption^[23,24]. Low education level was also associated with a higher risk of periodontitis which has also been reported by de Miguel-Infante *et al*^[13] and Yamamoto *et al*^[25]. People with lower education may have less exposure to information about oral health and less understanding about

the risk of periodontitis with T2DM. Unsurprisingly, less frequent brushing was associated with a higher risk of disease which has also been reported by Hong *et al*^[14] and Chang *et al*^[26]. Brushing removes food residue and reduces the reproduction of bacteria in the mouth, thereby decreasing the likelihood of developing periodontitis^[27].

The random forest model is widely used, and some studies have used it to predict the risk of nephropathy, peripheral neuropathy, and foot ulcers in patients with T2DM^[9,28,29]. We found that the random forest model was significantly better than the logistic regression model for the validation but not the test dataset. Both models had good predictive value with AUCs of 0.946 and 0.915, respectively. The random forest model is an ensemble learning method based on decision trees. Its advantage is that it requires less data, and its modeling process is more convenient and faster than logistic regression. However, the sensitivity, specificity, accuracy, recall, precision, and AUC of the training dataset of the random forest model in this study reached 1, indicating that the model may have been overfitting. A logistic regression model is a commonly used probability prediction model, which is simple to use and has strong predictive ability. Its advantage lies in that it can quantify the risk of disease through the OR value of variables, but it cannot intuitively determine the importance of each independent variable to the model prediction.

This study has some limitations: (1) This was a retrospective study based on the data of patients diagnosed with T2DM at our hospital. The study subjects and sources are from one site and the sample size included in the model is small, so there is a certain sample bias; (2) This study analyzed only a subset of the factors that may influence periodontitis; and (3) The random forest model may have an issue with overfitting. Future studies should include samples from other regions and a more comprehensive analysis of factors that influence periodontitis, and construct a more complete prediction model.

CONCLUSION

In conclusion, the factors influencing periodontitis in patients with T2DM were identified using logistic regression analysis. In patients with T2DM, the greater the age and HbA1c,

TC, and TG levels, the higher the risk of periodontitis. Our predictive models had good predictive value and could effectively predict the risk of periodontitis in patients with T2DM. The random forest and logistic regression prediction models can complement each other and provide a full analysis of the risk of disease and the importance of specific factors. In clinical practice, the results of this study can provide reference for the identification and intervention of early periodontitis in patients with T2DM.

ARTICLE HIGHLIGHTS

Research background

Periodontitis is a complication of type 2 diabetes mellitus (T2DM). With lifestyle changes and the acceleration of the aging process, the prevalence of periodontitis and diabetes is increasing annually.

Research motivation

Periodontitis can lead to tooth loosening and loss, decline in oral function, and reduced living standards.

Research objectives

This study aimed to explore and analyze the factors influencing periodontal disease in patients with T2DM, and construct prediction models for the risk of periodontal disease in patients with T2DM.

Research methods

We conducted a retrospective study in patients with T2DM hospitalized in our hospital to analyze the factors influencing periodontitis in patients with T2DM. We used random forest and logistic regression prediction models to assess the risk of specific factors in periodontitis.

Research results

This study found that the factors influencing periodontal disease in patients with T2DM were age, brushing frequency, education level, and glycosylated hemoglobin, total cholesterol, and triglyceride levels. The prediction models both had good predictive value.

Research conclusions

In this study, a random forest model was established and compared to a logistic regression model. The results showed that the random forest and logistic regression models had good predictive value and can accurately predict the risk of periodontitis in patients with T2DM.

Research perspectives

In the future, we will expand the sample size, combine samples from multiple regions, and include additional influencing factors to build a more complete prediction model.

REFERENCES

- 1 Yan Y, Wu T, Zhang M, Li C, Liu Q, Li F. Prevalence, awareness and control of type 2 diabetes mellitus and risk factors in Chinese elderly population. *BMC Public Health* 2022; **22**: 1382 [PMID: 35854279 DOI: 10.1186/s12889-022-13759-9]
- 2 Baeza M, Morales A, Cisterna C, Cavalla F, Jara G, Isamitt Y, Pino P, Gamonal J. Effect of periodontal treatment in patients with periodontitis and diabetes: systematic review and meta-analysis. *J Appl Oral Sci* 2020; **28**: e20190248 [PMID: 31939522 DOI: 10.1590/1678-7757-2019-0248]
- 3 Preshaw PM, Alba AL, Herrera D, Jepsen S, Konstantinidis A, Makrilakis K, Taylor R. Periodontitis and diabetes: a two-way relationship. *Diabetologia* 2012; **55**: 21-31 [PMID: 22057194 DOI: 10.1007/s00125-011-2342-y]
- 4 Stöhr J, Barbaresko J, Neuenschwander M, Schlesinger S. Bidirectional association between periodontal disease and diabetes mellitus: a systematic review and meta-

analysis of cohort studies. *Sci Rep* 2021; **11**: 13686 [PMID: 34211029 DOI: 10.1038/s41598-021-93062-6]

5 **Genco RJ**, Borgnakke WS. Diabetes as a potential risk for periodontitis: association studies. *Periodontol 2000* 2020; **83**: 40-45 [PMID: 32385881 DOI: 10.1111/prd.12270]

6 **Preshaw PM**, Bissett SM. Periodontitis: oral complication of diabetes. *Endocrinol Metab Clin North Am* 2013; **42**: 849-867 [PMID: 24286953 DOI: 10.1016/j.ecl.2013.05.012]

7 **Nitta H**, Katagiri S, Nagasawa T, Izumi Y, Ishikawa I, Izumiyama H, Uchimura I, Kanazawa M, Chiba H, Matsuo A, Utsunomiya K, Tanabe H, Takei I, Asanami S, Kajio H, Ono T, Hayashi Y, Ueki K, Tsuji M, Kurachi Y, Yamanouchi T, Ichinokawa Y, Inokuchi T, Fukui A, Miyazaki S, Miyauchi T, Kawahara R, Ogiuchi H, Yoshioka N, Negishi J, Mori M, Mogi K, Saito Y, Tanzawa H, Nishikawa T, Takada N, Nanjo K, Morita N, Nakamura N, Kanamura N, Makino H, Nishimura F, Kobayashi K, Higuchi Y, Sakata T, Yanagisawa S, Tei C, Ando Y, Hanada N, Inoue S. The number of microvascular complications is associated with an increased risk for severity of periodontitis in type 2 diabetes patients: Results of a multicenter hospital-based cross-sectional study. *J Diabetes Investig* 2017; **8**: 677-686 [PMID: 28129466 DOI: 10.1111/jdi.12633]

8 **Mikami R**, Mizutani K, Matsuyama Y, Gohda T, Gotoh H, Aoyama N, Matsuura T, Kido D, Takeda K, Saito N, Fujiwara T, Izumi Y, Iwata T. Association of type 2 diabetes with periodontitis and tooth loss in patients undergoing hemodialysis. *PLoS One* 2022; **17**: e0267494 [PMID: 35522619 DOI: 10.1371/journal.pone.0267494]

9 **Zou Y**, Zhao L, Zhang J, Wang Y, Wu Y, Ren H, Wang T, Zhang R, Wang J, Zhao Y, Qin C, Xu H, Li L, Chai Z, Cooper ME, Tong N, Liu F. Development and internal validation of machine learning algorithms for end-stage renal disease risk prediction model of people with type 2 diabetes mellitus and diabetic kidney disease. *Ren Fail* 2022; **44**: 562-570 [PMID: 35373711 DOI: 10.1080/0886022X.2022.2056053]

10 **Bian Y**, Liu C, Fu Z. Application value of combination therapy of periodontal curettage and root planing on moderate-to-severe chronic periodontitis in patients with type 2 diabetes. *Head Face Med* 2021; **17**: 12 [PMID: 33832490 DOI: 10.1186/s13005-020-00253-z]

- 11 **Barutta F**, Bellini S, Durazzo M, Gruden G. Novel Insight into the Mechanisms of the Bidirectional Relationship between Diabetes and Periodontitis. *Biomedicines* 2022; **10** [PMID: 35052857 DOI: 10.3390/biomedicines10010178]
- 12 **Romano F**, Perotto S, Mohamed SEO, Giraudi M, Bernardi S, Durazzo M, Gruden G, Aimetti M. Type 2 diabetes mellitus and periodontitis: Are diabetic patients aware about this bidirectional association? *Acta Diabetol* 2021; **58**: 1277-1280 [PMID: 33987719 DOI: 10.1007/s00592-021-01734-6]
- 13 **de Miguel-Infante A**, Martinez-Huedo MA, Mora-Zamorano E, Hernández-Barrera V, Jiménez-Trujillo I, de Burgos-Lunar C, Cardenas Valladolid J, Jiménez-García R, Lopez-de-Andrés A. Periodontal disease in adults with diabetes, prevalence and risk factors. Results of an observational study. *Int J Clin Pract* 2018: e13294 [PMID: 30444571 DOI: 10.1111/ijcp.13294]
- 14 **Hong M**, Kim HY, Seok H, Yeo CD, Kim YS, Song JY, Lee YB, Lee DH, Lee JI, Lee TK, Ahn HS, Ko YH, Jeong SC, Chae HS, Sohn TS. Prevalence and risk factors of periodontitis among adults with or without diabetes mellitus. *Korean J Intern Med* 2016; **31**: 910-919 [PMID: 27604799 DOI: 10.3904/kjim.2016.031]
- 15 **Eke PI**, Wei L, Borgnakke WS, Thornton-Evans G, Zhang X, Lu H, McGuire LC, Genco RJ. Periodontitis prevalence in adults ≥ 65 years of age, in the USA. *Periodontol 2000* 2016; **72**: 76-95 [PMID: 27501492 DOI: 10.1111/prd.12145]
- 16 **Wu CZ**, Yuan YH, Liu HH, Li SS, Zhang BW, Chen W, An ZJ, Chen SY, Wu YZ, Han B, Li CJ, Li LJ. Epidemiologic relationship between periodontitis and type 2 diabetes mellitus. *BMC Oral Health* 2020; **20**: 204 [PMID: 32652980 DOI: 10.1186/s12903-020-01180-w]
- 17 **Qureshi A**, Haque Z, Bokhari SAH, Baloch AA. Evaluation of HbA1c in type-2 diabetes mellitus patients with periodontitis: preliminary findings of three-arm clinical trial. *J Pak Med Assoc* 2020; **70**: 1350-1356 [PMID: 32794485 DOI: 10.5455/JPMA.22016]
- 18 **Dhir S**, Wangnoo S, Kumar V. Impact of Glycemic Levels in Type 2 Diabetes on Periodontitis. *Indian J Endocrinol Metab* 2018; **22**: 672-677 [PMID: 30294579 DOI: 10.4103/ijem.IJEM_566_17]

- 19 **Simpson TC**, Clarkson JE, Worthington HV, MacDonald L, Weldon JC, Needleman I, Iheozor-Ejiofor Z, Wild SH, Qureshi A, Walker A, Patel VA, Boyers D, Twigg J. Treatment of periodontitis for glycaemic control in people with diabetes mellitus. *Cochrane Database Syst Rev* 2022; **4**: CD004714 [PMID: 35420698 DOI: 10.1002/14651858.CD004714.pub4]
- 20 **Sanz M**, Ceriello A, Buysschaert M, Chapple I, Demmer RT, Graziani F, Herrera D, Jepsen S, Lione L, Madianos P, Mathur M, Montanya E, Shapira L, Tonetti M, Vegh D. Scientific evidence on the links between periodontal diseases and diabetes: Consensus report and guidelines of the joint workshop on periodontal diseases and diabetes by the International Diabetes Federation and the European Federation of Periodontology. *J Clin Periodontol* 2018; **45**: 138-149 [PMID: 29280174 DOI: 10.1111/jcpe.12808]
- 21 **Persson GR**. Dental geriatrics and periodontitis. *Periodontol 2000* 2017; **74**: 102-115 [PMID: 28429479 DOI: 10.1111/prd.12192]
- 22 **Ding C**, Du F, Li L, Chen Y. Synergistic effect of blood lipids and uric acid on periodontitis in patients with type 2 diabetes. *Am J Transl Res* 2023; **15**: 1430-1437 [PMID: 36915719]
- 23 **Jiménez-Corona M**, Falcón-Flores J, Borges-Yáñez A, Castrejón-Pérez R, Jiménez-Corona A. Dyslipidemia and severe periodontitis among patients with type 2 diabetes. *Salud Publica Mex* 2021; **63**: 331-332 [PMID: 34098619 DOI: 10.21149/11890]
- 24 **Thomas B**, Prasad RB, Shetty S, Vishakh R. Comparative Evaluation of the Lipid Profile in the Serum of Patients with Type II Diabetes Mellitus and Healthy Individuals with Periodontitis. *Contemp Clin Dent* 2017; **8**: 96-101 [PMID: 28566858 DOI: 10.4103/ccd.ccd_1160_16]
- 25 **Yamamoto T**, Tanaka M, Kuribayashi N, Okuguchi F, Isotani H, Iwamoto M, Sugimoto H, Nakagawa O, Minabe M, Fuchida S, Mochida Y, Yokoyama H. Low education is associated with poor periodontal status in patients with type 2 diabetes mellitus: A cross-sectional study. *Clin Exp Dent Res* 2021; **7**: 419-428 [PMID: 33258300 DOI: 10.1002/cre2.363]
- 26 **Chang Y**, Lee JS, Lee KJ, Woo HG, Song TJ. Improved oral hygiene is associated with decreased risk of new-onset diabetes: a nationwide population-based cohort study. *Diabetologia* 2020; **63**: 924-933 [PMID: 32128623 DOI: 10.1007/s00125-020-05112-9]

- 27 **Leite RS**, Marlow NM, Fernandes JK, Hermayer K. Oral health and type 2 diabetes. *Am J Med Sci* 2013; **345**: 271-273 [PMID: 23531957 DOI: 10.1097/MAJ.0b013e31828bdebf]
- 28 **Wang J**, Xue T, Li H, Guo S. Nomogram Prediction for the Risk of Diabetic Foot in Patients With Type 2 Diabetes Mellitus. *Front Endocrinol (Lausanne)* 2022; **13**: 890057 [PMID: 35909507 DOI: 10.3389/fendo.2022.890057]
- 29 **Wu B**, Niu Z, Hu F. Study on Risk Factors of Peripheral Neuropathy in Type 2 Diabetes Mellitus and Establishment of Prediction Model. *Diabetes Metab J* 2021; **45**: 526-538 [PMID: 34352988 DOI: 10.4093/dmj.2020.0100]

Footnotes

Institutional review board statement: The study was reviewed and approved by the First People's Hospital of Wenling (approval No. KY-2023-2035-01).

Informed consent statement: Informed consent was waived due to the retrospective nature of this study.

Conflict-of-interest statement: All the authors report no relevant conflicts of interest for this article.

Data sharing statement: The datasets used in this study can be obtained from the corresponding author upon reasonable request.

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Figure Legends

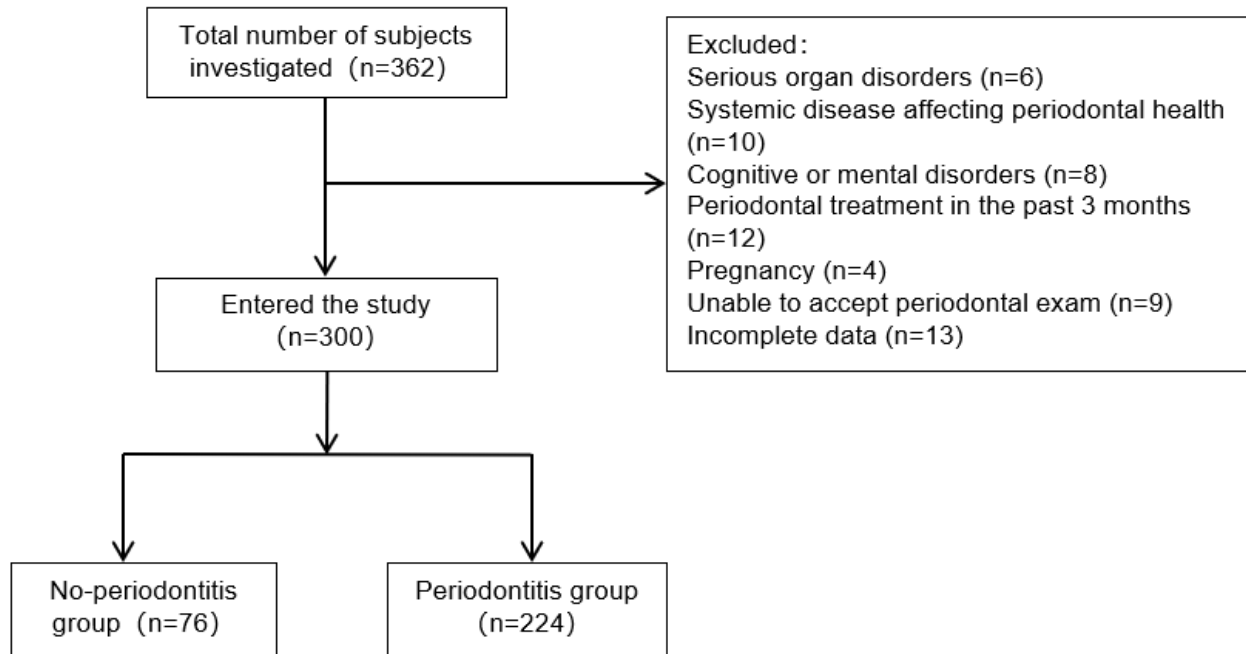


Figure 1 Flow diagram.

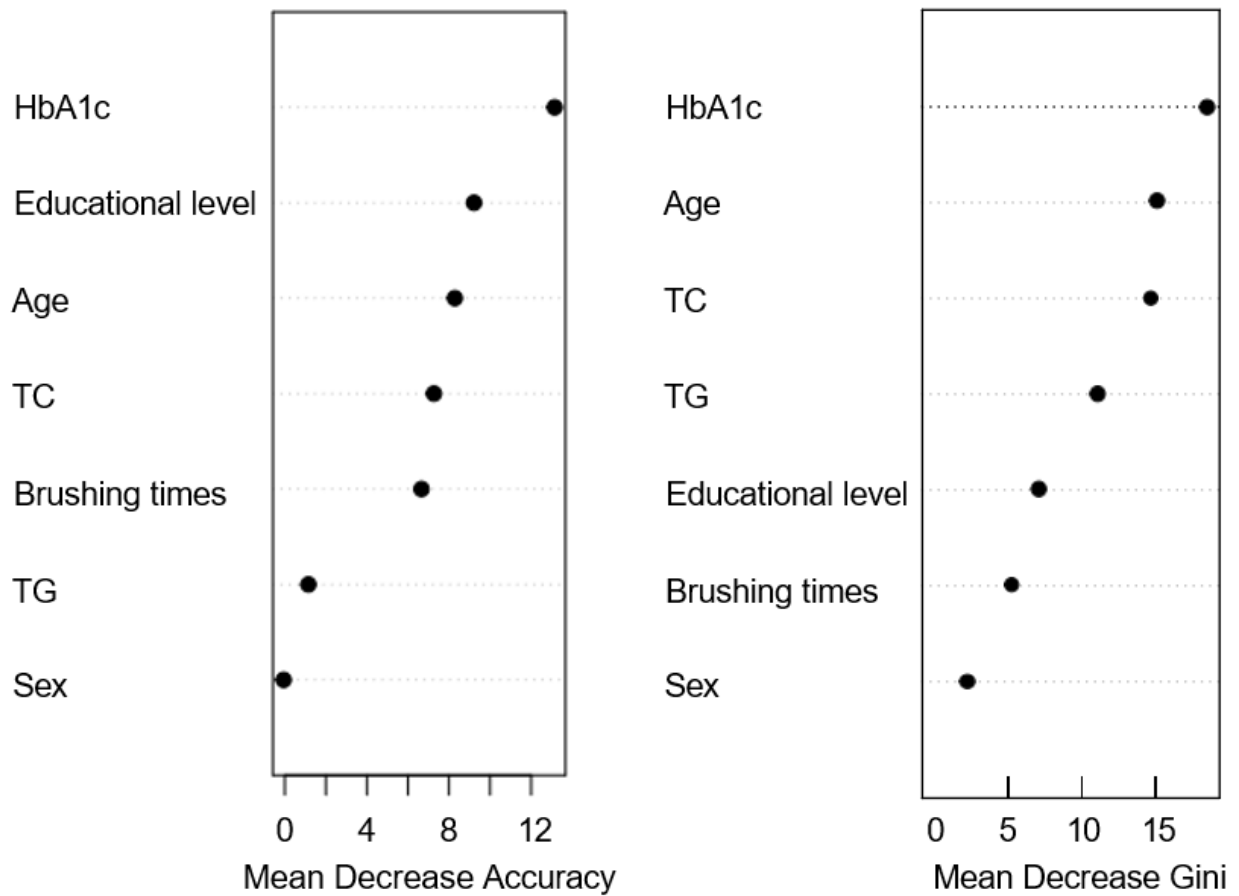


Figure 2 Importance of variables in the random forest model. HbA1c: Glycosylated hemoglobin; TC: Total cholesterol; TG: Triglyceride.

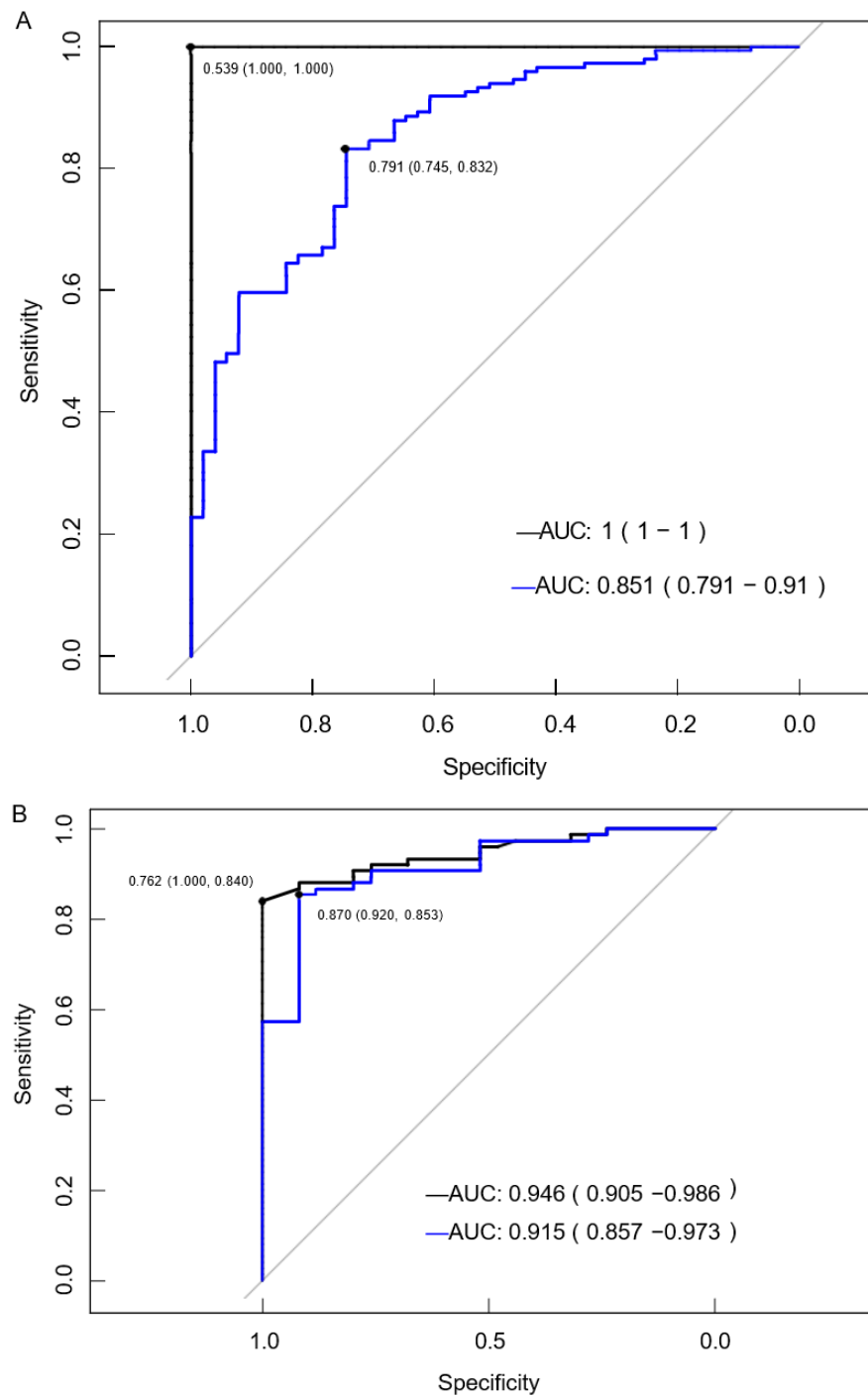


Figure 3 Receiver operating characteristic curves of the random forest (black) and logistic regression (blue) models. A: For the training set; B: For the validation set. AUC: Area under curve.

Table 1 General information

Factor	Total number (<i>n</i> = 300)	Periodontitis group (<i>n</i> = 224)	Non- periodontitis group (<i>n</i> = 76)	<i>t</i> / χ^2 / <i>Z</i>	<i>P</i> value
Sex, <i>n</i> (%)				4.050	0.044
Male	164 (54.67)	130 (58.04)	34 (44.74)		
Female	136 (45.33)	94 (41.96)	42 (55.26)		
Age (yr), (mean \pm SD)	59.18 \pm 11.40	60.91 \pm 9.49	54.11 \pm 14.69	-4.646	< 0.001
Education level <i>n</i> (%)				-3.626	< 0.001
Below high school education	118 (39.33)	98 (43.75)	20 (26.31)		
High school education	98 (32.67)	76 (33.93)	22 (28.95)		
Above high school education	84 (28.00)	50 (22.32)	34 (44.74)		
Monthly income, <i>n</i> (%)				-1.890	0.059
< 2000 yuan	65 (21.67)	54 (24.11)	11 (14.47)		
2000-5000 yuan	133 (44.33)	99 (44.20)	34 (44.74)		
> 5000 yuan	102 (34.00)	71 (31.70)	31 (40.79)		

Table 2 Lifestyle factors

Factor	Total (<i>n</i> = 300)	Periodontitis group (<i>n</i> = 224)	Non-periodontitis group (<i>n</i> = 76)	χ^2	<i>P</i> value
Smoking, <i>n</i> (%)				3.500	0.061
No	238 (79.33)	172 (76.79)	66 (86.84)		
Yes	62 (20.67)	52 (23.21)	10 (13.16)		
Alcohol consumption, <i>n</i> (%)				0.460	0.497
No	249 (83.00)	184 (82.14)	65 (85.53)		
Yes	51 (17.00)	40 (17.86)	11 (14.47)		
Regular exercise, <i>n</i> (%)					
No	142 (47.33)	109 (48.66)	33 (43.42)	0.625	0.429
Yes	158 (52.67)	115 (51.34)	43 (56.58)		
Brushing frequency, <i>n</i> (%)				29.248	< 0.001
≥ 2 times/d	137 (45.67)	82 (36.61)	55 (72.37)		
< 2 times/d	163 (54.33)	142 (63.39)	21 (27.63)		

Table 3 Physical health status

Factor	Total number (<i>n</i> = 300)	Periodontitis group (<i>n</i> = 224)	Non-periodontitis group (<i>n</i> = 76)	<i>t</i> / <i>Z</i>	<i>P</i> value
BMI (kg/m ²), median (P25, P75)	24.96 (22.36, 26.81)	24.86 (22.13, 26.65)	25.61 (23.26, 27.23)	-1.223	0.221
TG (mmol/L)	1.51 ± 0.28	1.53 ± 0.28	1.44 ± 0.26	-2.678	0.008
< 1.7	226 (75.33)	163 (72.77)	63 (82.89)		
≥ 1.7	74 (26.67)	61 (27.23)	13 (17.11)		
TC (mmol/L)	4.72 ± 0.77	4.86 ± 0.78	4.33 ± 0.59	-5.361	< 0.001
< 5.18	212 (70.67)	144 (64.29)	68 (89.47)		
≥ 5.18	88 (29.33)	80 (35.71)	8 (10.53)		
HDL-C (mmol/L)	1.09 ± 0.19	1.08 ± 0.20	1.12 ± 0.15	1.777	0.077
< 1.04, <i>n</i> (%)	114 (38.00)	93 (41.52)	21 (27.63)		
≥ 1.04, <i>n</i> (%)	186 (62.00)	131 (58.48)	55 (72.37)		
LDL-C (mmol/L)	2.42 ± 0.29	2.44 ± 0.29	2.39 ± 0.30	-1.183	0.238
< 3.37, <i>n</i> (%)	300 (100.00)	224 (100.00)	76 (100.00)		
≥ 3.37, <i>n</i> (%)	0 (0.00)	0 (0.00)	0 (0.00)		
HbA1c (mmol/L)	7.86 ± 1.17	8.10 ± 1.16	7.14 ± 0.85	-6.596	< 0.001
< 6.5%, <i>n</i> (%)	37 (12.33)	17 (7.59)	20 (26.32)		
6.5%-7.5%, <i>n</i> (%)	83 (27.66)	52 (23.21)	31 (40.79)		
7.5%-8.5%, <i>n</i> (%)	90 (30.00)	71 (31.70)	19 (25.00)		
≥ 8.5%, <i>n</i> (%)	90 (30.00)	84 (37.50)	6 (7.89)		

BMI: Body mass index; TG: Triglyceride; TC: Total cholesterol; HDL-C: High-density lipoprotein cholesterol; LDL-C: Low-density lipoprotein cholesterol; HbA1c: Glycosylated hemoglobin.

Table 4 Assignment and description of related factors affecting the occurrence of periodontitis in patients with type 2 diabetes mellitus

Factor	Assignment of value
Sex	Female = 0 Male = 1
Age	Continuous variables
Brushing frequency	≥ 2 times/d = 0 < 2 times/d = 1
Education level	Below high school education = 0 High school education = 1 Above high school education = 2
HbA1c	Continuous variables
TC	Continuous variables
TG	Continuous variables

HbA1c: Glycosylated hemoglobin; TC: Total cholesterol; TG: Triglyceride.

Table 5 Multivariate logistic regression analysis of occurrence of periodontitis in patients with type 2 diabetes mellitus

Factor	β	SE	Wald χ^2	P value	OR (95%CI)
Age	0.046	0.015	9.813	0.002	1.047 (1.017-1.078)
Sex	0.622	0.346	3.230	0.072	1.863 (0.945-3.674)
Brushing frequency	1.459	0.353	17.073	< 0.001	4.303 (2.154-8.599)
Education level	-0.639	0.212	9.065	0.003	0.528 (0.348-0.800)
HbA1c	0.934	0.185	25.392	< 0.001	2.545 (1.770-3.661)
TC	1.055	0.260	16.451	< 0.001	2.872 (1.725-4.781)
TG	1.196	0.600	3.967	0.046	3.306 (1.019-10.723)

HbA1c: Glycosylated hemoglobin; TC: Total cholesterol; TG: Triglyceride; OR: Odds ratio; 95%CI: 95% confidence interval.

Table 6 Efficacy of the two models in predicting the occurrence of periodontitis in patients with type 2 diabetes mellitus in the training dataset

Model	Sensitivity	Specificity	Accuracy	Recall	Precision	AUC (95%CI)
Random forest	1.000	1.000	1.000	1.000	1.000	1.000 (1.000-1.000)
Logistic regression	0.569	0.919	0.830	0.569	0.707	0.851 (0.791-0.910)

AUC: Area under curve; CI: Confidence interval.

Table 7 Efficacy of the two models in predicting the occurrence of periodontitis in patients with type 2 diabetes mellitus in the validation dataset

Model	Sensitivity	Specificity	Accuracy	Recall	Precision	AUC (95%CI)
Random forest	0.520	0.960	0.850	0.520	0.813	0.946 (0.905-0.986)
Logistic regression	0.640	0.907	0.840	0.640	0.696	0.915 (0.857-0.973)

AUC: Area under curve; CI: Confidence interval.