

ORIGINAL RESEARCH

Torus Palatinus in Taiwan Patients Receiving Peritoneal Dialysis and Hemodialysis: A Prospective Observational Study

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Purpose: A consensus about the pathogenesis of torus palatinus (TP) in patients receiving dialysis still eludes the scientific community. This prospective observational study investigated the epidemiology of TP in peritoneal dialysis and hemodialysis patients and analyzed the influences of multiple pathogenic factors such as mineral and bone disorders, genetic, environmental or nutritional triggers, progression of age, heredity, climatologic or biomechanical causes, and hyperparathyroidism on the formation of TP.

Methods: Between 2013 and 2016, a total of 575 chronic dialysis patients (441 on hemodialysis and 134 on peritoneal dialysis) were recruited from Chang Gung Memorial Hospital, Taiwan. Patients were stratified into two groups based on the presence (n = 179) or absence (n = 396) of TP. Demographic, oral examination, laboratory, and dialysis data were collected for analysis. Student's t-test was used to analyze the quantitative variables and Chisquare or Fisher's exact test for categorical variables. Univariate binary logistic regression analysis was conducted to determine the predictors for TP and multivariate binary logistic regression analysis to identify significant associated factors.

Results: The prevalence of TP in dialysis patients in this study was 31.1% (28.3% for hemodialysis and 40.3% for peritoneal dialysis). Patients with TP were younger (54.6 \pm 13.4 versus 58.9 ± 14.7 years, P = 0.001) and mostly female (60.3 versus 41.2%, P < 0.001). Most TP cases (55.3%) were small in size (<2 cm), with the flat shape (56.4%) being the most common followed by the spindle (17.9%), nodular (17.3%), and lobular (8.4%) shapes. A longer duration of dialysis was associated with TP ≥2 cm than with TP <2 cm (94.4 ± 85.9 versus 72.8 ± 59.1 months, P = 0.048). Multivariate logistic regression revealed that female gender (odds ratio 2.108, 95% confidence interval 1.455–3.055, P < 0.001) and younger age (odds ratio 0.982; 95% confidence interval 0.969-0.994, P = 0.005) were significant predictors for TP.

Conclusion: The prevalence of TP in chronic dialysis patients is 31.1%, higher in patients receiving peritoneal dialysis (40.3%) than hemodialysis (28.3%). Female gender and younger age are significant predictors associated with TP.

Keywords: end-stage renal disease, hemodialysis, peritoneal dialysis, torus palatinus

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Introduction

Torus palatinus (TP) is non-pathological bony deposition, which gradually develops along the midline of the palate and consists of cortical and trabecular bones. TP is often discovered incidentally after middle age due to the lack of symptoms that can be detected during routine dental examinations. Surgical removal of TP may be necessary when patients present the following: speech and masticatory disturbances, Chang et al Dovepress

trauma and ulceration of the mucosa from wearing the dental prosthesis,² prosthetic instability, intubation difficulties,³ obstructive sleep apnea,⁴ cancer phobia, or as TP becomes the potential source of autogenous cortical bone for grafting.

A consensus about the pathogenesis of TP in patients receiving dialysis still eludes the scientific community. Many associated factors have been proposed, such as: mineral and bone disorders;5 genetic;6 environmental or nutritional triggers; progression of age; heredity; 8 climatologic⁹ or biomechanical (functional) causes. The prevalence of treated end-stage renal disease in Taiwan has been one of the highest in the world. 10 Chronic renal failure alters bone metabolism through multiple mechanisms. For example, phosphate retention and decreased conversion of vitamin D lead to hypocalcemia and stimulate parathyroid chief cells to produce more parathyroid hormone. Padbury et al,11 have found decreased cortical density and increased likelihood of oral tori in patients with primary hyperparathyroidism. Long-term dialysis also inhibits bone homeostasis as a result of aluminum deposition that interferes with bone mineralization.¹²

In terms of prevalence, Sisman et al¹³ reported a TP prevalence of 41.7% in peritoneal dialysis patients. There were also studies reporting markedly different prevalence rates in different ethnic groups.^{6,14-16} According to Chiang et al,¹⁷ the prevalence of TP in the general population of Taiwan was 21.1%. Also in Taiwan, the prevalence of TP in hemodialysis patients was 23.5% as found by Chao et al⁵ and 28.9% reported by Tai et al,¹⁸ respectively, and 34.3% in peritoneal dialysis patients according to Hsu et al.¹⁹

These studies^{5,18,19} also concluded that neither hyperparathyroidism nor inflammation-malnutrition syndrome contributes to the risk of developing tori in dialysis patients. Meanwhile, Sisman et al¹³ found evidence for the association between the peritoneal dialysis duration and the TP size. Tai et al¹⁸ demonstrated that younger age, female gender, higher blood concentration of phosphate, and lower blood concentration of bicarbonate were significant predictors for TP in hemodialysis patients.

Therefore, the objective of this prospective observational study was to investigate the epidemiology of TP in peritoneal dialysis and hemodialysis patients, and analyzed the influences of multiple pathogenic factors such as mineral and bone disorders, genetic, environmental or nutritional triggers, progression of age, heredity, climatologic or biomechanical causes, hyperparathyroidism on the formation of TP.

Patients and Methods

Ethical Statement

This clinical study followed the Declaration of Helsinki for Human Experimentation and was approved by the Medical Ethics Committee of the Chang Gung Memorial Hospital. The Institutional Review Board numbers assigned to the study were 102-2761B and 104-6913C.

Sample Size

Sample size was determined using G*Power software version 3.1.9.7. An odds ratio of female gender to predict TP was 2.305 according to a pilot study. Therefore, the sample size as calculated by G*Power with odds ratio of 2.305, alpha error of 0.05 and power of 0.95 was 255.

Study Design and Setting

Between 2013 and 2016, a total of 575 chronic dialysis patients (441 on hemodialysis and 134 on peritoneal dialysis) were recruited from two branches of the Chang Gung Memorial Hospital located in Linkou and Taoyuan. All primary data were collected according to procedures outlined in strengthening the reporting of observational studies in epidemiology (STROBE).

Inclusion and Exclusion Criteria

All dialysis patients who agreed to participate were eligible for inclusion in this study. Patients undergoing hemodialysis or peritoneal dialysis for less than 6 months, having been hospitalized, operated on or received kidney transplant in the preceding 3 months as well as those with cancer were excluded from the study.

Consenting Process

All patients provided written informed consent prior to participating in this study.

Measures – Dialysis, Diagnosis of TP and Molar Relationship

Dialysis. Hemodialysis patients underwent treatment 3 times a week. Each treatment lasted approximately 4 hours, using a single-use hollow-fiber artificial kidney with modified cellulose-based polyamide or polysulfone membranes. The dialysate used was a standard ionic composition and bicarbonate-based buffer. Peritoneal dialysis prescriptions were based on the peritoneal membrane characteristics determined by the peritoneal equilibration test. Intermittent therapy was prescribed to patients with high membrane transport,

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Figure I Torus palatinus. Intraoral view of the edentulous hemodialysis patient showed a flat torus palatinus (asterisk) along the midline of the hard palate.

and continuous therapy, to those with average or low membrane transport.

Diagnosis of TP. TP was diagnosed through clinical inspection and palpation (Figure 1). The size of TP was determined by the maximum elevation of the outgrowth of tori. TPs were characterized as ≥ 2 cm or < 2 cm using a periodontal probe, as described by Gorsky et al.²⁰ The shapes of TP were grouped as flat, spindle, nodular, or lobular according to the criteria suggested by Jainkittivong et al.²¹

Molar relationship. The molar relationship was assessed and scored as none, Class I, II or III, based on Angle's classification.²²

Statistical Analysis

All data were tested for normality of distribution and equality of standard deviation before analysis. Student's t-test was used to analyze the quantitative variables, and Chi-square or Fisher's exact test, for categorical variables.

Univariate binary logistic regression analysis was conducted to determine the predictors for TP, and multivariate binary logistic regression analysis, to identify significant associated factors. A P value of less than 0.05 was selected as the significance threshold to reject the null hypothesis. All analyses were performed using IBM SPSS Statistics Version 20.0.

Results

Table 1 presents the baseline characteristics of participating patients. The prevalence of TP was 31.1% (28.3% for hemodialysis and 40.3% for peritoneal dialysis). Patients with TP were younger (54.6 \pm 13.4 versus 58.9 \pm 14.7 years, P = 0.001) and mostly female (60.3 versus 41.2%, P < 0.001).

As outlined in Table 2, the majority of TPs (55.3%) were of small size (< 2 cm). The flat-shaped TP (56.4%) was the most common, followed by the spindle- (17.9%), nodular- (17.3%), and lobular-shaped (8.4%) TPs.

The molar relationship could not be identified in 280 (48.7%) patients due to the loss of their first permanent

Table 2 Patterns of TP in Dialysis Patients (n = 179)

Variable	
Size ≥ 2cm, n (%)	80 (44.7)
< 2cm, n (%)	99 (55.3)
Shape	
Flat, n (%)	101 (56.4)
Spindle, n (%)	32 (17.9)
Nodular, n (%)	31 (17.3)
Lobular, n (%)	15 (8.4)

Abbreviation: TP, torus palatinus.

Table I Baseline Demographics of Patients Receiving Dialysis (n = 575)

Variable	All Patients (n = 575)	Patients with TP (n = 179)	Patients Without TP (n = 396)	P value
Age, year	57.6 ± 14.4	54.6 ±13.4	58.9 ± 14.7	0.001**
Female gender, n (%)	271 (47.1)	108 (60.3)	163 (41.2)	< 0.001***
Dialysis duration, months	89.2 ± 74.5	82.5 ± 72.9	92.3 ± 75.1	0.143
Type of dialysis				0.009**
Hemodialysis, n (%)	441 (76.7)	125 (69.8)	316 (79.8)	
Peritoneal dialysis, n (%)	134 (23.3)	54 (30.2)	80 (20.2)	
Hypertension, n (%)	311 (54.1)	101 (56.4)	210 (53.0)	0.449
Diabetes mellitus, n (%)	185 (32.2)	55 (30.7)	130 (32.8)	0.617
Alcohol consumption, n (%)	32 (5.6)	8 (4.5)	24 (6.1)	0.441
Smoking habit, n (%)	69 (12.0)	17 (9.5)	52 (13.1)	0.214

Notes: **P < 0.01; ***P < 0.001. Abbreviation: TP, torus palatinus. Chang et al Dovepress

Table 3 Molar Relationships in Patients Receiving Dialysis (n = 575)

Variable	All Patients (n=575)	Patients with TP (n=179)	Patients Without TP (n=396)	P value
No, n (%) Class I, n (%)	280 (48.7) 204 (35.5)	80 (44.7) 75 (14.9)	200 (50.5) 129 (32.6)	0.184
Class II, n (%) Class III, n (%)	49 (8.5) 42 (7.0)	13 (7.3) 11 (6.1)	36 (9.1) 31 (7.8)	

Abbreviation: TP, torus palatinus.

molars (Table 3). There was no significant difference between the two study groups (P = 0.184).

Patients with TP had higher blood levels of phosphate $(5.5\pm1.3~{\rm versus}~5.1\pm1.4~{\rm mg/dL}, P=0.010)$ and bicarbonate $(21.2\pm3.7~{\rm versus}~21.1\pm3.4~{\rm mmol/L}, P=0.020)$, and lower residual renal function $(4.4\pm1.3~{\rm versus}~4.6\pm1.4~{\rm mL/min/1.73~m^2}, P=0.012)$ than patients without TP (Table 4).

Although patients with TP had lower blood concentrations of the intact parathyroid hormone than patients without TP, the difference was not significant (328.9 \pm 340.4 versus 359.6 \pm 364.2 pg/mL, P=0.341). No significant differences in inflammatory variables such as high sensitivity C-reactive protein (7.6 \pm 11.8 versus 8.7 \pm 16.8mg/L, P=0.433) and nutritional variables such as albumin (3.9 \pm 0.3 versus 3.9 \pm 0.4 g/dL, P=0.220) were seen between the two groups. There were also no significant differences in dialysis parameters between the two groups, such as Kt/V (1.9 \pm 0.4 versus 1.8 \pm 0.4 g/dL, P=0.071) and normalized protein catabolic rate (1.1 \pm 0.5 versus 1.1 \pm 0.4 g/dL, P=0.470).

The multivariate logistic regression analysis revealed that female gender (odds ratio 2.108, 95% confidence interval 1.455–3.055, P < 0.001) and younger age (odds ratio 0.982; 95% confidence interval 0.969–0.994, P = 0.005) were significant predictors for TP (Table 5).

Table 4 Laboratory Findings and Dialysis Parameters in Patients Receiving Dialysis (n = 575)

Variables	All Patients (n = 575)	Patients with TP (n = 179)	Patients Without TP (n = 396)	P value
Blood urea nitrogen, mg/dL	67.5 ± 19.1	67.9 ± 17.7	67.3 ± 19.6	0.722
Creatinine, mg/dL	11.0 ± 2.7	11.3 ± 2.9	10.8 ± 2.6	0.092
Uric acid, mg/dL	6.8 ± 1.3	7.0 ± 1.3	6.8 ± 1.3	0.068
Sodium, mEq/L	137.5 ± 3.2	137.5 ± 3.2	137.5 ± 3.2	0.847
Potassium, mEq/L	4.5 ± 0.8	4.6 ± 0.8	4.5 ± 0.7	0.517
Calcium, mg/dL	9.6 ± 1.0	9.6 ± 0.9	9.6 ± 1.0	0.746
Phosphate, mg/dL	5.2 ± 1.3	5.5 ± 1.3	5.1 ± 1.4	0.010*
Bicarbonate, mmol/L	21.7 ± 3.5	21.2 ± 3.7	21.1 ± 3.4	0.020*
Fasting glucose, mg/dL	116.0 ± 54.9	114.0 ± 51.2	117.8 ± 56.4	0.446
Albumin, g/dL	3.9 ± 0.3	3.9 ± 0.3	3.9 ± 0.4	0.220
Alkaline phosphatase, U/L	82.3 ± 57.9	80.0 ± 48.2	84.3 ± 61.7	0.229
Total cholesterol, mg/dL	168.5 ± 39.7	176.7 ± 39.0	164.8 ± 39.4	0.001**
High-density lipoprotein, mg/dL	43.3 ± 14.8	42.8 ± 13.9	43.5 ± 15.2	0.561
Low-density lipoprotein, mg/dL	111.4 ± 71.6	117.3 ± 67.6	108.7 ± 73.3	0.180
Triglyceride, mg/dL	156.2 ± 132.3	170.2 ± 140.5	149.8 ± 128.0	0.085
White blood cell count, 10 ³ /uL	6.7 ± 2.2	7.1 ± 2.3	6.6 ± 2.1	0.030*
Red blood cell count, 106/uL	3.6 ± 0.6	3.5 ± 0.5	3.6 ± 0.6	0.313
Hematocrit, %	31.5 ± 3.9	31.2 ± 3.9	31.6 ± 3.9	0.369
Mean corpuscular volume, fL	88.9 ± 7.3	89.9 ± 6.3	88.9 ± 7.8	0.962
Platelet count, 10 ³ /uL	195.8 ± 69.5	211.3 ± 71.2	188.8 ± 67.6	< 0.001***
Iron, ug/dL	71.5 ± 45.9	76.3 ± 68.1	69.83 ± 31.0	0.092
Total iron binding capacity, ug/dL	261.9 ± 54.1	262.9 ± 54.9	260.6 ± 53.9	0.635
Ferritin, ng/mL	332.3 ± 390.5	314.4 ± 266.2	340.3 ± 435.3	0.463
Residual renal function, mL/min/1.73 m ²	4.6 ± 1.4	4.4 ± 1.3	4.6 ± 1.4	0.012*
Intact parathyroid hormone, pg/mL	350.0 ± 357.0	328.9 ± 340.4	359.6 ± 364.2	0.341
High sensitivity C-reactive protein, mg/L	8.3 ± 15.4	7.6 ± 11.8	8.7 ± 16.8	0.433
Kt/V	1.9 ± 0.4	1.9 ± 0.4	1.8 ± 0.4	0.071
Normalized protein catabolic rate, g/kg/day	1.1 ± 0.5	1.1 ± 0.5	1.1 ± 0.4	0.470

Notes: *P < 0.05; **P < 0.01; ***P < 0.001.

Abbreviation: TP, torus palatinus.

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Table 5 Logistic Regression Analysis of Risk Factors for TP (n = 575)

Variable	Univariate Analysis			Multivariate Analysis		
	Odds Ratio	95% Confidence Interval	P value	Odds Ratio	95% Confidence Interval	P value
Age (each increase of I year)	0.980	0.968-0.992	0.001**	0.982	0.969–0.994	0.005**
Bicarbonate (each increase of I mmol/L)	0.945	0.900-0.993	0.026*	0.966	0.917-1.017	0.186
Residual renal function (each increase of 1 mL/min/1.73 m ²)	0.832	0.720-0.962	0.013*	0.921	0.796-1.066	0.269
Female gender (yes)	2.174	1.517–3.117	< 0.001***	2.108	1.455–3.055	< 0.001***
Phosphate (each increase of 1 mg/dL)	1.188	1.042-1.356	0.010*	1.094	0.945-1.266	0.230

Notes: P < 0.05; P < 0.01; P < 0.01.

Abbreviation: TP, torus palatinus.

As shown in Table 6, a longer duration of dialysis is associated with TP \geq 2 cm than with TP \leq 2 cm (94.4 \pm 85.9 versus 72.8 \pm 59.1 months, P = 0.048).

Discussion

This study examined the largest series of TPs in the current dialysis literature. The prevalence of TP in chronic dialysis patients is 31.1%, higher in patients receiving peritoneal dialysis (40.3%) than hemodialysis (28.3%). Notably, female gender and younger age are significant predictors for TP in dialysis patients.

The mechanism for TP formation induced by female gender remains unclear. Nevertheless, the observed female predominance of TP may be related to genetic, environmental or functional factors. First, TP is transmitted as an autosomal dominant trait, and it is believed that there may be a dominant type linked to the X chromosome. 6 Second, the different prevalence rates between females and males might be attributed to the same environmental factors, but exerted differently to women and men. Furthermore, differences in food consumption, oral pathology, and extramasticatory dental activities could also produce a differential spreading of TPs.²³ In Eskimo inhabitants, women exhibit TPs more frequently than men because they repeatedly use teeth for hide preparation. The condition is common in Eskimos who place great pressures on

Table 6 Comparison of Dialysis Duration Between Patients with $TP \ge 2$ Cm and TP < 2 Cm (n = 179)

Variable	Patients with TP ≥2 cm (n = 80)	Patients with TP <2 cm (n = 99)	P value
Duration of dialysis, months	94.4 ± 85.9	72.8 ± 59.1	0.048*

Note: *P < 0.05.

Abbreviation: TP, torus palatinus.

their masticatory system and upper jaw. These forces are medially directed towards the peak of the palatine vault, and creating thickening of the palatal roof along the median suture to limit this pressure.²⁴ Dietary habits and nutritional disturbances are also considered by some researchers as potential causes.²⁵ Eggen and Natvig suggested that the consumption of saltwater fish containing omega 3 unsaturated fatty acids and vitamin D could possibly encourage bone growth and increase the chances of TPs.²⁶ Many previous studies have also found a higher prevalence of TP in females than males,^{9,27,28} as in the current study where 60.3% of patients with TP were females.

Similarly, the mechanism for TP formation induced by younger age also remains uncertain. Nevertheless, the more prevalence of TP in the younger age may be associated with nutritional and functional factor. First, a decreasing ability to chew and absorb food is likely in elderly people, which may result in malnutrition and, thereby, could possibly affect the occurrence of TP.²⁹ Second, as progression of age there may be decrease in numbers of teeth as result to extraction of teeth and periodontal disease, this result in decrease of masticatory function as the persons become partially or completely edentulous, so less occlusal stress get and it can be consider as functional factors. In terms of the link between age and TP, the onset of tori can be found as early as in the prenatal period, newborns, and young children who have not been subjected to mechanical stimuli. 30 Afterwards, tori tend to grow with age, possibly due to the continuous and increased functional occlusal stress from teeth to the underlying alveolar bone. 31 Eggen et al also pointed out that TP seemed to be a dynamic phenomenon capable of bone growth and resorption remodeling. 14 TP is usually discovered incidentally after middle age. ^{1,4} In this study, patients with TP aged 54.6 ± 13.4 years. This finding was similar to those from previous Chang et al Dovepress

studies.^{32–34} However, the prevalence of tori decreases especially after the age of 50. This drop in prevalence might be attributable to malnutrition^{32,34} and a decrease in the masticatory function and less occlusal stress³⁵ due to loss of teeth from caries or periodontal diseases as the patient ages. Oral tori also subside in size after the fifth decade of life because of the reduced number of teeth as a result of bone resorption.³⁶ In the current study, the average age of patients with TP was younger than those without (P = 0.001). This finding echoed that tori are more frequently observed during the middle age than at older ages.^{14,27}

Different prevalence rates of TP have been reported for different population groups. ^{14,15} Chiang et al ¹⁷ found the TP prevalence to be 21.1% in the general population of Taiwan. In dialysis patients, Chao et al ⁵ and Tai et al ¹⁸ reported TP prevalence rates of 23.5% and 28.9%, respectively, among hemodialysis patients, and Hsu et al, ¹⁹ a TP prevalence rate of 34.3% among peritoneal dialysis patients. In the current study, the prevalence of TP was 31.1% - 28.3% for hemodialysis and 40.3% for peritoneal dialysis.

A higher blood concentration of phosphate was also found in patients with TP than without (P = 0.010) in the current study. The different TP prevalence rates between the general population and chronic dialysis patients thus might be attributed to the existence of underlying disorders. Reduced renal function produces hypocalcemia by favoring phosphate retention. Low serum calcium leads to a compensatory parathyroid hyperactivity, resulting in elevated phosphate excretion, decrease in calcium excretion, and increased removal of calcium from bones by osteoclast activation.³⁷ Hyperphosphatemia elevates the blood concentration of fibroblast growth factor 23, reduction in active vitamin D synthesis, and the tendency toward hypocalcemia - all are potent stimuli for secondary hyperparathyroidism. ^{38–40} Hyperparathyroidism depletes cortical bone and stimulate formation of cancellous bone, resulting in changes in the contour of the affected bony structure. The preferential loss of cortical bone and increased formation of trabecular bone, particularly in the oral cavity, are usually the early signs of renal osteodystrophy. 11

Our study also found that the number of patients with TP < 2 cm (55.3%) was greater than that with TP \geq 2 cm (44.7%), which mirrors the study results by Jainkittivong et al, ²⁸ Reichart et al⁹ and Sisman et al. ¹³ Furthermore, a longer duration of dialysis was associated with TP \geq 2 cm than with TP < 2 cm (P = 0.048). This resonates with the observations by Sisman et al ¹³ where a significant relationship existed

between TP size and duration of peritoneal dialysis. They went on to suggest underlying disorders such as renal osteo-dystrophy as the likely culprit.

TP is a benign disease and most TP manifestations in this study are small in size. Many of our dialysis patients were often unaware of their presence before this study. Therefore, there is no major clinical significance of TP in dialysis patients. Nevertheless, the clinical observations of female and younger age predominance of TP are interesting, and warranted more researches in this area. Finally, the limitations of this study included small patient population, short follow up duration, and lacking pathological evaluation.

Conclusions

The prevalence of TP in chronic dialysis patients in Taiwan is 31.1%, higher in patients receiving peritoneal dialysis (40.3%) than hemodialysis (28.3%). Patients with TP are younger than patients without TP, and are mostly female. Most TP manifestations are small in size. The flat-shaped TP is the most common, followed by the spindle-, nodular-, and lobular-shaped. A longer duration of dialysis is associated with TP \geq 2 cm than with TP < 2 cm. The female gender and younger age are significant predictors for TP in chronic dialysis patients.

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Disclosure

The authors declare no conflicts of interest in this work.

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