#### **ORIGINAL ARTICLE**

# Interproximal implant thread exposure after initial bone remodeling as a risk indicator for peri-implantitis

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#### Abstract

Background: Due to the clinical challenges involved in successfully treating peri-implantitis, it is imperative to identify patient- and implant-level risk factors for its prevention. The main goal of this retrospective longitudinal radiographic and clinical study was to investigate whether interproximal radiographic implant thread exposure after physiological bone remodeling may be a risk factor for peri-implantitis. The secondary goal was to evaluate several other potential risk indicators.

Methods: Of 4325 active dental school patients having implants placed, 165 partially edentulous adults (77 men, 88 women) aged 30–91 with  $\geq$ 2 years of follow-up upon implant restoration were included. Implants with  $\geq 1$  interproximal thread exposed (no bone-to-implant contact) (n = 98, 35%) constituted the test group and those without exposed threads (n = 182, 65%) the control group. Descriptive, binary, and multivariate regression analyses were evaluated for goodness of fit. Wald tests were used to evaluate for significance set at 0.05.

Results: Of the 280 implants (98 test, 182 control), 8 (2.9%) failed over a mean follow-up period of 7.67 (±2.63) years, and 27 implants (19 test, 8 control) developed peri-implantitis, with the exposed group having eight-fold (7.82 times) adjusted greater odds than the non-exposed. The risk increased four-fold (3.77 times) with each thread exposed. No other patient- or implant-related potentially confounding risk factors were identified.

Conclusions: Exposed interproximal implant threads after physiologic bone remodeling may be an independent risk indicator for incident peri-implantitis. Hence, clinicians should closely monitor patients with implant threads that have no bone-to-implant contact for incident peri-implantitis.

#### **KEYWORDS**

bone resorption, dental implants, periodontics, radiography, tooth loss

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# **1** | INTRODUCTION

Peri-implantitis (PI) is defined as an inflammatory lesion in the tissues surrounding the implant with progressing of bone loss beyond the expected physiologic bone remodeling.<sup>1,2</sup> PI is the most common complication in implant dentistry,<sup>3,4</sup> affecting around 20% of patients<sup>5–7</sup> and 13% of implants,<sup>6,7</sup> with study results ranging widely.

Because successful treatment of PI is so challenging and the outcome unpredictable,<sup>4,8</sup> it is imperative to prevent PI from developing, which necessitates identification of its local and systemic risk factors<sup>3</sup> for potential mitigation.

According to the 2017 World Workshop on the Classification of Periodontal and Peri-implant Diseases and Conditions ("2017 World Workshop"), a history of periodontitis, poor plaque control, and lack of regular maintenance therapy might be considered risk indicators of PI; however, other factors such as smoking, diabetes, width of keratinized tissue, titanium particles, and prosthesis design need to be further evaluated.<sup>1</sup>

It is currently accepted that PI is caused by bacterial challenge in a susceptible host,<sup>9</sup> possibly in combination with a foreign body immune reaction.<sup>10</sup>

Several studies have focused on the roles of the patient (plaque control and compliance with professional maintenance visits) and of the provider (non-surgical or surgical therapies and maintenance) in the development of PI.<sup>8,11-18</sup> Implant design has been discussed extensively regarding osseointegration, but few studies have explored its role in disease onset,<sup>19,20</sup> so the role of the implant topography in PI requires further investigation.<sup>21</sup> Implant topography can be categorized as macro- and microdesign, respectively. The macrodesign pertains to the shape of the implant body as well as the design and number of threads and is an established key factor for osseointegration as a crucial element for primary implant stability and possibly for bone-toimplant contact (BIC).<sup>22-24</sup> Implant macrodesign has also been hypothesized to be a possible factor contributing to peri-implant disease.<sup>21,25–27</sup> In support of this hypothesis, greater PI prevalence was found in implants with triple thread, with a microthreaded collar, and with a cylindric shape.27

The microdesign concerns the chemically or mechanically treated implant surface, such as by acid etching, sandblasting, titanium plasma spraying, and hydroxyapatite coating.<sup>28–30</sup> Moderately rough implant surfaces were associated with lower prevalence rates of PI,<sup>7</sup> but due to the limited quality of evidence on the topic, more studies are necessary to evaluate the relationship between implant microdesign and PI.<sup>31</sup> As a potential risk for PI,<sup>32</sup> bone graft was also recorded.<sup>32</sup>

A clinical study observed that small bony buccal dehiscence defects developed greater-than-expected vertical bone loss 6 months after implant placement.<sup>33</sup> However, no study has explored the impact of the interproximal thread exposure on the development of PI.

Thus, the main aim of this retrospective longitudinal study was to investigate whether radiological interproximal implant thread exposure after physiological bone remodeling may be a potential risk indicator for incident PI. The secondary goal was to identify other potential patient- or implant-related risk factors for incident PI.

#### 2 | MATERIALS AND METHODS

The study protocol was approved by the University of Michigan Medical School Institutional Review Board (Study #HUM00194509) and was conducted in accordance with the Helsinki Declaration adopted in 1964 and 1975,<sup>34,35</sup> as revised in 2013.<sup>36</sup> This retrospective investigation included implants placed and restored by graduate students or faculty at the University of Michigan School of Dentistry between January 2000 and September 2017. Eligible participants needed to fulfill the following inclusion criteria: 1) partially edentulous area restored with  $\geq 1$ implant with a documented follow-up period of  $\geq 2$  years after implant loading; 2) clinical data and high-quality periapical radiographs available at the time of implant placement (T0), prosthetic restoration (T1), 1 year after prosthetic restoration (T2, radiograph exposed at that time as per institutional protocol), and at follow-up of  $\geq 2$  years after prosthetic restoration (T3); 3) available information about the implant brand as well as the surface micro- and macrostructure; 4) presence of opposing teeth/restored implants (occlusion); 5) no active periodontitis at the time of implant placement (T0). Exclusion criteria were a) presence of PI in the test group at T2; b) potentially confounding comorbidities, such as a history of uncontrolled diabetes mellitus, radiation or chemotherapy, psychologic or psychiatric issues; and c) receipt of treatment or maintenance visits external to the study institution. Physical and digital records for potentially eligible patients were screened and evaluated by four examiners (A.S., M.Q., M.S., and L.W.) who subsequently extracted the data. Any disagreement that arose during the screening for eligibility and the data collection process was resolved through discussion with the principal investigator (A.R.).

# 2.1 | Data collection and classification

Relevant patient information was extracted, including age at the time of implant placement (T0), sex, smoking habit ( $\geq 1$  cigarette/day), diabetes mellitus (validated via the patient's medical records), history of periodontitis,



and number of maintenance appointments. A positive history of periodontitis was determined following the case definition for periodontitis proposed by the 2017 World Workshop<sup>37</sup> based on periodontal charts and radiographs. Detailed implant specific data collected included the number of implants and their positions (location in the edentulous jaw area), implant design (bone or soft tissue level), brand, length, diameter, neck design, retention type of restoration (cement or screw), and splinting. Bone grafting (yes/no) was recorded, and the type of implant-abutment connection and neck designs were also collected. Moreover, data were collected on the distance between threads (pitch) and the implant macrosurface, such as thread designs (buttress, reverse buttress, square, progressive square, and V-shaped), which are schematically illustrated in Figure 1.<sup>24</sup> Details about the microsurface recorded included type of surface (microtextured and sandblasted, large-grit, acid-etched). The implants were divided into four different categories according to their roughness (S<sub>a</sub>): smooth (S<sub>a</sub> < 0.5  $\mu$ m); minimally rough (S<sub>a</sub> 0.5–1.0  $\mu$ m), moderately rough (S<sub>a</sub> > 1.0–2.0  $\mu$ m), and rough (S<sub>a</sub> > 2.0  $\mu$ m). <sup>38,39</sup>

Implants were divided by radiographic evaluation of interproximal (mesial/distal) BIC 1 year after prosthetic restoration (T2): 1) absence of BIC with  $\geq$ 1 proximal implant thread (test group, "exposed") and 2) no thread without BIC (control group, "non-exposed"). A thread was regarded radiographically exposed when the adjacent bone did not completely cover its surface.<sup>40</sup> Exposed and non-exposed implant threads are illustrated conceptually in

Figure 2 and radiographically in Figures S1 and S2 (in online *Journal of Periodontology*).

# 2.2 | Definition of outcomes

Based on our predefined outcomes, data analysis for implant failure, prevalence of PI, marginal bone loss, and numbers of threads exposed was performed. Two distinct follow-up periods were defined prior to data acquisition: a) follow-up to assess implant survival and b) follow-up to assess occurrence of PI, marginal bone loss, and number of interproximal (mesial or distal) threads exposed (with no BIC). The follow-up duration based on implant survival was defined as the time between implant placement (T0) and T4, defined as the last visit, during which each implant was classified as present or explanted. The followup based on the occurrence of PI, marginal bone loss, and number of threads exposed was defined as the duration of time between T2 and exposure of the last radiograph on which peri-implant bone could be clearly visualized (T3). The time between T2 and T3 is referred to as the "radiograph period." In case of concomitancy between T3 and T4 (the last X-rays available and the last patient visit), the two follow-up durations were identical.

Implant failure was defined as a removed, lost, mobile, or fractured implant.<sup>41</sup> PI was defined as proposed by the 2017 World Workshop<sup>2</sup> and was used to classify cases in a binary fashion as either positive (1) or negative (0) for PI. Because baseline data were available, a PI diagnosis JOURNAL OF



**FIGURE 2** Development of marginal bone loss leading to exposed implant thread (no bone-to-implant contact). Implant placed at bone level (T1) (**A**). Bone loss after remodeling 1 year after implant prosthetic restoration (T2) (**B**). Close-up from panel B showing the most coronal implant thread exposed (**C**). (Conceptual model not showing any prosthetic restoration). (Please also see radiographs from study patients with and without interproximal thread exposure in Figures S1 and S2 in the online *Journal of Periodontology*)

was based on 1) progressive bone loss beyond initial bone remodeling, 2) increased probing depth, and 3) presence of bleeding and/or suppuration on gentle probing. Marginal bone level (MBL) was defined as the distance between the most coronal portion of the implant expected to present radiographic bone contact (for tissue-level implants, the interface between the polished collar and rough surface, and for bone-level implants, the platform level) to the most coronal point of the implant body in contact with bone. The MBL and the count of the exposed threads at T2 and T3 were radiographically assessed by two authors (A.R., M.S.) at the mesial and distal aspects of the affected implants using commercially available image software.\* If significant differences arose (>0.5 mm for bone loss and >1 thread for the thread count), a third reviewer (H.L.W.) was included for reassessing the radiographs in a joint session to reach a final judgment. Repeated measurements of 15 implants were initially conducted to quantify mean interexaminer agreement measurement errors for MBL, which was  $0.32 (\pm 0.2)$  mm.

# 2.3 | Statistical analysis

The statistical analysis included descriptive analyses of categorical (absolute and relative frequencies) and continuous (mean, standard deviation, range, and median) variables for the total sample and stratified by study group (exposed/non-exposed threads) using dedicated statistical software.<sup>†</sup> The outcome PI diagnosis (yes/no) was related to all independent variables using multilevel binary logistic regression with generalized estimation equations (GEE). Raw odds ratios (OR) and 95% confidence intervals (CI) were obtained from the Wald chi-square statistic.

Then, multivariate models were applied to adjust by potential confounding factors. The goodness of fit of different GEE estimates (for different matrix correlations) was assessed by QIC (quasi-likelihood under the independence model criterion) statistic. Significance level in all analyses was set to 5% ( $\alpha = 0.05$ ). A post hoc power analysis was conducted. A sample size of 280 independent implants would provide 90.9% power with a confidence level of 95% to detect an OR of 3 as significant, using logistic regression models. Since the implants were not independent due to the two-level (patient and implant) data structure, this power needed correction. With each patient providing 1.75 implants on average and assuming a within-subject correlation of 0.5 (moderate), the correcting coefficient (D) was 1.35. Therefore, 280 dependent implants provide the same power as 207 independent implants, estimated at 80.4% under the mentioned conditions.

# 3 | RESULTS

# 3.1 | Clinical characteristics and demographic profiles

Records from a total of 4325 active patients who had received implant therapy at the University of Michigan School of Dentistry were screened for potential inclusion. A total of 1287 patients were excluded due to <2 years postimplant restoration follow-up period, 2423 patients due to absence of  $\geq 1$  radiograph or periodontal chart, 352 patients due to lack of information about brand and other implant characteristics, 53 patients due to presence of fixed full-arch restorations, and 45 due to ambiguous or incomplete charts. Hence, 165 patients were included in the study, including 77 males (46.7%) and 88 females (53.3%) with a mean age of 62.5 ( $\pm$ 11.7) years ranging from 30 to 91 years at baseline (T0). A total of 280 implants were

<sup>&</sup>lt;sup>\*</sup> ImageJ, US National Institutes of Health, Bethesda, Maryland.

<sup>&</sup>lt;sup>†</sup> SPSS, Chicago, Illinois.

included (n = 98 in the test group; n = 182 in the control group). Characteristics of the sample at patient and implant levels are displayed in Table 1.

# 3.2 | PI and marginal bone loss

Overall, the PI rate was 9.6% (27/280) in the total sample of implants. About one-fifth (19.4%) of the implants in the test group and 4.4% in the control group developed PI. Results from simple binary logistic regression using GEE (Table 2) show that an increasing number of threads exposed and the square thread design significantly increased the probability of developing PI. Moreover, increasing patient age significantly decreased this probability. No other confounder obtained statistically significant effect in the bivariate analyses.

A multivariate model (Table 3) considering these findings and adjusting for potential confounders (duration of and mean annual number of maintenance visits during the radiographic period [T2 to T3]) showed that thread exposure remained a significant factor for increasing the likelihood of PI, with the risk of PI increasing almost eight-fold with each additional exposed thread (OR 7.82; 95% CI, 1.91–32.03; p = 0.004). Splinting was also associated with greater risk for PI (OR 3.49; 95% CI, 1.02–12.05; p = 0.047). Each year of increased age was associated with 5% lower risk of a PI diagnosis (OR 0.95; 95% CI, 0.92–0.99; p = 0.016).

No association was found between PI and any other implant macro- or microsurface design nor a history of periodontitis. The mean annual crestal bone loss between T2 to T3 was 0.26 ( $\pm$ 0.65) mm in the exposed (test group) versus 0.11 ( $\pm$ 0.31) mm per year in the non-exposed (control) group (p = 0.05). Each additional exposed thread significantly increased the odds of PI almost four-fold (OR 3.77; 95% CI, 1.82–7.82; p < 0.001) (Figure 3A; see also Table S1 in online *Journal of Periodontology*).

# 3.3 | Implant failure

Each group lost four implants. The failure rate was at 2.9% (8/280) in the total sample (4.1% in the test group and 2.2% in the control group), a statistically non-significant difference (p = 0.470) (see Table S2 in online *Journal of Periodontology*). The probability of failure increased with the number of exposed threads, with each additional thread increasing the probability of failure about three times (OR 3.13; 95% CI, 1.01–9.66; p < 0.001) (Figure 3B; see also Table S3 in online *Journal of Periodontology*). Other than older age (OR 0.97; 95% CI, 0.94–1.00; p = 0.049),

there were no other variables identified to potentially prevent implant failure.

# 4 | DISCUSSION

Because PI is difficult to arrest once established, identification of its modifiable risk factors is key for prevention. In implant treatment planning, execution, and maintenance, all possible measures to prevent development of exposed threads must be taken. Indeed, the results demonstrated an eight-fold increased risk for PI in implants with exposed threads compared to those with non-exposed threads. The risk increased four-fold with each additional thread exposed, and splinting was associated with 3.49 times greater risk for incident PI, whereas no other confounding patient-level factor (except for age) or implant macro- or microdesign feature was identified.

The reasons for exploring other potential risk factors were to not only identify them but to ensure statistically that such confounders might not actually be causing the incident PI instead of the thread exposure. Successful treatment of PI is very demanding. Retaining such success through maintenance proved to be challenging as well as shown by a systematic review and meta-analysis, where there was merely <5% reduction in the risk of implant loss for patients undergoing periodic maintenance therapy compared to those who did not.<sup>42</sup> In a recent study, patients without maintenance therapy had 4.25 times greater risk for PI.<sup>16</sup> Nonetheless, in the present study, the mean number of annual maintenance visits was found to not be associated with incident PI.

Splinting was found to present a 3.49 times greater risk for PI in multivariate analyses adjusted for duration and mean annual number of maintenance visits (Table 3). This finding is in contrast to the conclusions of a systematic review that a) there was no difference in MBL between splinted and non-splinted implant restorations<sup>43</sup> and b) splinting was associated with lower risk for implant failure.<sup>43</sup> On the contrary, our finding was in agreement with another study that also found greater risk of PI in splinted individual implant restorations, although threeunit bridges supported by two implants had significantly less risk for PI.<sup>44</sup> It should be noted that our study was not able to assess the accessibility for cleaning the implants and their restorations.

Our findings suggest that apart from splinting, the only modifiable statistically significant patient- and implantrelated risk factor for incident PI was the number of implant threads exposed 1 year after prosthetic implant restorations, and the latter impact was dose-dependent. To the best of our knowledge, this is the first time such 

 TABLE 1
 Patient- and implant-level characteristics of the implants placed in the 165 patients (N = 280 implants)

	Total, mean	Non-exposed (0 threads exposed), mean $\pm$ SD or	Exposed ( $\geq 1$ thread exposed), mean $\pm$ SD or
Characteristic	± SD or <i>n</i> (%)	n (%)	n (%)
Number of implants	280	182 (65.0)	98 (35.0)
Patient age at T0, years	$63.0 \pm 11.3$	$62.7 \pm 11.1$	$63.3 \pm 11.5$
Sex			
Male	123 (43.9)	76 (41.8)	47 (48.0)
Female	157 (56.1)	106 (58.2)	51 (52.0)
Smoking (≥1 cigarette/day)			
No	241 (86.1)	161 (88.5)	80 (81.6)
Yes	39 (13.9)	21 (11.5)	18 (18.4)
Diabetes			
No	245 (87.5)	155 (85.2)	90 (91.8)
Yes	35 (12.5)	27 (14.8)	8 (8.2)
History of periodontitis			
No	185 (66.1)	122 (67.0)	63 (64.3)
Yes	95 (33.9)	60 (33.0)	35 (35.7)
Duration of follow-up period			
T0–T1, months	$8.81 \pm 4.72$	$8.41 \pm 4.57$	9.55 ± 4.94
T2–T3 (radiograph period), years	$4.60 \pm 2.52$	$4.51 \pm 2.66$	4.78 ± 2.25
T0–T4, years	$7.67 \pm 2.63$	$7.53 \pm 2.45$	$7.91 \pm 2.93$
Edentulous site			
Incisor/canine	20 (7.2)	12 (6.6)	8 (8.2)
Premolar	110 (39.3)	70 (38.5)	40 (40.8)
Molar	150 (53.6)	100 (54.9)	50 (51.0)
Arch			
Maxilla	99 (35.4)	65 (35.7)	34 (34.7)
Mandible	181 (64.6)	117 (64.3)	64 (65.3)
Bone graft			
No	212 (76.0)	138 (76.2)	74 (75.5)
Yes	67 (24.0)	43 (23.8)	24 (24.5)
Implant surface			
MTX	105 (37.5)	87 (47.8)	18 (18.4)
TiUnite	103 (36.8)	32 (17.6)	71 (72.4)
SLA	43 (15.4)	42 (23.1)	1 (1.0)
SLA active	2 (0.7)	2 (1.1)	0
Friadent plus	7 (2.5)	7 (3.8)	0
Nanotite	9 (3.2)	6 (3.3)	3 (3.1)
RBT	10 (3.6)	6 (3.3)	4 (4.1)
СМІ	1 (0.4)	0 (0.0)	1 (1.0)
Roughness (S <sub>a</sub> )			
Smooth/minimally rough (S <sub>a</sub> $\leq$ 1.0 $\mu$ m)	7 (2.5)	7 (3.8)	0
Moderate ( $S_a > 1.0-2.0 \ \mu m$ )	170 (60.7)	143 (78.6)	27 (27.6)
Rough ( $S_a > 2.0 \mu m$ )	103 (36.8)	32 (17.6)	71 (72.4)

(Continues)

## TABLE 1 (Continued)

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	Total, mean	Non-exposed (0 threads exposed), mean $\pm$ SD or	Exposed ( $\geq 1$ thread exposed), mean $\pm$ SD or
Characteristic	$\pm$ SD or <i>n</i> (%)	n (%)	n (%)
Connection			
Internal hexagon	124 (44.4)	99 (54.4)	25 (25.8)
External hexagon	52 (18.6)	8 (4.4)	44 (45.4)
Morse taper	45 (16.1)	44 (24.2)	1 (1.0)
Internal hexagon with Morse taper	20 (7.2)	12 (6.6)	8 (8.2)
Internal trilobe	31 (11.1)	12 (6.6)	19 (19.6)
Morse taper cone connection	7 (2.5)	7 (3.8)	0
Neck design			
0.5 machined collar (Zimmer)	25 (9.0)	17 (9.3)	8 (8.2)
0.5 MTX collar	67 (24.0)	58 (31.9)	9 (9.3)
1.0 machined collar (Zimmer)	13 (4.7)	12 (6.6)	1 (1.0)
Fine micron feature	9 (3.2)	6 (3.3)	3 (3.1)
Laser-Lok collar	10 (3.6)	6 (3.3)	4 (4.1)
Misc. machined collar (Nobel)	22 (7.9)	8 (4.4)	14 (14.4)
Microrough shoulder	7 (2.5)	7 (3.8)	0
Microthreads	29 (10.4)	16 (8.8)	13 (13.4)
Smooth collar	44 (15.8)	43 (23.6)	1 (1.0)
Threaded	53 (19.0)	9 (4.9)	44 (45.4)
Thread design			
Buttress	46 (16.4)	44 (24.2)	2 (2.0)
Progressive square	7 (2.5)	7 (3.8)	0
Reverse buttress	93 (33.2)	26 (14.3)	67 (68.4)
Square	20 (7.1)	12 (6.6)	8 (8.2)
V-shaped	114 (40.7)	93 (51.1)	21 (21.4)
Implant level			
Bone level	197 (70.6)	110 (60.4)	87 (89.7)
Tissue level	82 (29.4)	72 (39.6)	10 (10.3)
Length			
<11 mm	79 (28.3)	52 (28.6)	27 (27.8)
11–12 mm	131 (47.0)	88 (48.4)	43 (44.3)
>12 mm	69 (24.7)	42 (23.1)	27 (27.8)
Diameter			
<4 mm	52 (22.4)	34 (20.0)	18 (29.0)
4–4.5 mm	81 (34.9)	63 (37.1)	18 (29.0)
>4.5 mm	99 (42.7)	73 (42.9)	26 (41.9)
Retention			
Cemented	201 (72.0)	134 (73.6)	67 (69.1)
Screwed	75 (26.9)	45 (24.7)	30 (30.9)
Overdenture	3 (1.1)	3 (1.6)	0
Splinted			
No	204 (72.9)	144 (79.1)	60 (61.2)
Yes	76 (27.1)	38 (20.9)	38 (38.8)
			(Continues)

#### **TABLE 1** (Continued)

Characteristic	Total, mean ± SD or <i>n</i> (%)	Non-exposed (0 threads exposed), mean $\pm$ SD or n (%)	Exposed ( $\geq 1$ thread exposed), mean $\pm$ SD or n (%)
Number of annual maintenance visits during radiograph period (T2–T3)			
$\leq 1$	63 (23.1)	41 (22.8)	22 (23.7)
>1-≤2	104 (38.1)	73 (40.6)	31 (33.3)
>2-≤3	77 (28.2)	47 (26.1)	30 (32.3)
>3	29 (10.6)	19 (10.6)	10 (10.8)
Number of annual maintenance visits (T0–T4)			
≤0.5	61 (22.4)	43 (24.0)	18 (19.4)
>0.5-≤1	59 (21.7)	45 (25.1)	14 (15.1)
>1-≤1.5	91 (33.5)	54 (30.2)	37 (39.8)
>1.5	61 (22.4)	37 (20.7)	24 (25.8)

Abbreviations: MTX, microtextured; RBT, resorbable blast texturing; SLA, sandblasted, large-grit, acid-etched; T0, time of implant placement; T1, time of prosthetic restoration; T2, 1 year after prosthetic restoration; T3, time of exposure of last radiograph on which peri-implant bone could be clearly visualized; T4, time of last patient visit.



**FIGURE 3** Predicted probability of peri-implantitis (PI) (**A**) and of implant failure (**B**) by the number of exposed threads (N = 280 implants). Implant failure is defined as removed, lost, mobile, or fractured implant.<sup>41</sup> T2, 1 year after implant prosthetic restoration

conclusion has been demonstrated by rigorous research, even though this result seems intuitive. Since the body of literature appears to be void of relevant findings regarding the number of exposed threads, we cannot compare this main finding to prior research results.

Interestingly, severity of periodontitis was not a significant factor for incidence of PI, which is in accord with our group's earlier findings in another study population among patients at the same institution, where only periodontitis Grade C was associated with incident PI.<sup>15</sup> This finding is also in line with the results of the meta-analysis published in 2016, which obviously could not have applied the 2017 World Workshop case definitions for either disease.<sup>42</sup> A systematic review by Doornewaard and coworkers supports our findings that implant surface roughness was not a significant factor in PI.<sup>39</sup> It is noteworthy that we applied the current classification of both periodontitis and PI defined by the 2017 World Workshop, and therefore any direct comparison to prior research would benefit from reassessing the classification of both diseases in the older studies.

Despite the multitude of operators and potentially changing protocols related to implant placement and restoration at a dental school over a period of 18 years,

TABLE 2	Risks of incident peri-implantitis by patient, implant, and prosthesis characteristics during the total study period (T0–T4).
Results from u	inadjusted binary logistic regression analyses with generalized estimation equations ( $N = 280$ implants)

Characteristic	Total, mean $\pm$ SD	Peri-implantitis,	OR	95% CI	n value
Number of implants	280	$n(\pi)$	UK	95% CI	<i>p</i> value
Study group	200	27 (9.0)			
Non avnosad (0 throads avnosad)	182 (65.0)	9(4 4)	1		
Exposed (>1 thread exposed)	182(03.0)	3(4.4)	1 5 23	2 10-13 0	~0 001***
Patient age at T0, years	98(33.0)	19 (19.4)	0.05	2.10-13.0	0.001
Sev	$05.0 \pm 11.5$		0.95	0.92-0.99	0.008
Male	123 (43.0)	16 (13 0)	1		
Female	123(43.9) 157(561)	10(13.0)	0.50	0 18-1 40	0 190
Smoking (>1 cigarette/day)	157 (50.1)	11 (7.0)	0.50	0.10 1.40	0.170
No	241 (86 1)	26 (10.8)	1		
Yes	39 (13 9)	1(2.6)	0.22	0 03-1 77	0 154
Diabetes	57 (15.7)	1 (2.0)	0.22	0.05 1.77	0.154
No	245 (87 5)	23 (9 4)	1		
Yes	35 (12.5)	4 (11.4)	1.25	0.26-5.93	0.783
History of periodontitis		. ()	1120	0.20 0.00	01100
No	185 (66.1)	15(8.1)	1		
Yes	95 (33.9)	12 (12.6)	1.64	0.61-4.43	0.331
Duration of follow-up period		()			
T0–T1. months	8.81 + 4.72		1.05	0.93-1.18	0.458
T2–T3 (radiograph period), years	$4.60 \pm 2.52$		1.08	0.84–1.39	0.546
T0–T4, years	$7.67 \pm 2.63$		1.03	0.79–1.33	0.841
Edentulous site					0.552
Incisor/canine	20 (7.2)	1(5)	1		
Premolar	110 (39.3)	12 (10.9)	2.33	0.42-12.9	0.334
Molar	150 (53.6)	14 (9.3)	1.96	0.26-15.0	0.519
Arch					
Maxilla	99 (35.4)	9 (9.1)	1		
Mandible	181 (64.6)	18 (9.9)	1.10	0.38-3.21	0.856
Bone graft					
No	212 (76.0)	22 (10.4)	1		
Yes	67 (24.0)	5 (7.5)	0.70	0.23-2.13	0.525
Implant surface					0.194
MTX	105 (37.5)	6 (5.7)	1		
TiUnite	103 (36.8)	15 (14.6)	2.81	0.82–9.61	0.099
SLA	43 (15.4)	2 (4.7)	0.81	0.15-4.37	0.801
SLA active	2 (0.7)	0	n/a	n/a	n/a
Friadent plus	7 (2.5)	0	n/a	n/a	n/a
Nanotite	9 (3.2)	1 (11.1)	2.06	0.18-23.9	0.563
RBT	10 (3.6)	3 (30.0)	7.07	0.77-64.9	0.084
СМІ	1 (0.4)	0	n/a	n/a	n/a
Roughness (S <sub>a</sub> )					
Smooth/minimally rough (S <sub>a</sub> <1.0 $\mu$ m)	7 (2.5)	0	n/a	n/a	n/a
Moderate (S <sub>a</sub> 1.0–2.0 $\mu$ m)	170 (60.7)	12 (7.1)	1		
Rough (S <sub>a</sub> > 2.0 $\mu$ m)	103 (36.8)	15 (14.6)	2.24	0.82-6.13	0.115

(Continues)

# TABLE 2 (Continued)

	Total, mean $\pm$ SD	Peri-implantitis,			
Characteristic	or n (%)	n (%)	OR	95% CI	<i>p</i> value
Connection					0.275
Internal hexagon	124 (44.4)	10 (8.1)	1		
External hexagon	52 (18.6)	6 (11.5)	1.49	0.40-5.47	0.550
Morse taper	45 (16.1)	2 (4.4)	0.53	0.11-2.62	0.437
Internal hexagon with Morse taper	20 (7.2)	5 (25.0)	3.80	0.82–17.7	0.089
Internal trilobe	31 (11.1)	4 (12.9)	1.69	0.37-7.72	0.499
Morse taper cone connection	7 (2.5)	0	n/a	n/a	n/a
Neck design					0.308
0.5 machined collar (Zimmer)	25 (9.0)	3 (12.0)	1		
0.5 MTX collar	67 (24.0)	3 (4.5)	0.34	0.04–2.78	0.317
1.0 machined collar (Zimmer)	13 (4.7)	0	n/a	n/a	n/a
Fine micron feature	9 (3.2)	1 (11.1)	0.92	0.06-13.5	0.317
Laser-Lok collar	10 (3.6)	3 (30.0)	3.14	0.27-36.9	0.362
Machined collar (Zimmer)	22 (7.9)	2 (9.1)	0.73	0.10-5.62	0.765
Microrough shoulder	7 (2.5)	0	n/a	n/a	n/a
Microthreads	29 (10.4)	7 (24.1)	2.33	0.37-14.9	0.309
Smooth collar	44 (15.8)	2 (4.5)	0.35	0.05-2.65	0.309
Threaded	53 (19.0)	6 (11.3)	0.94	0.16-5.66	0.943
Thread design					0.080
Buttress	46 (16.4)	2 (4.3)	1		
Progressive square	7 (2.5)	0	n/a	n/a	n/a
Reverse buttress	93 (33.2)	13 (14.0)	3.58	0.77–16.6	0.105
Square	20 (7.1)	5 (25.0)	7.33	1.16-46.4	0.034*
V-shaped	114 (40.7)	7 (6.1)	1.44	0.28-7.39	0.663
Implant level					
Bone level	197 (70.6)	22 (11.2)	1		
Tissue level	82 (29.4)	5 (6.1)	0.52	0.16-1.69	0.274
Length					0.280
<11 mm	79 (28.3)	5 (6.3)	1		
11–12 mm	131 (47.0)	17 (13.0)	2.21	0.76-6.41	0.146
>12 mm	69 (24.7)	5 (7.2)	1.16	0.29-4.67	0.838
Diameter					0.978
<4 mm	52 (22.4)	4 (7.7)	1		
4–4.5 mm	81 (34.9)	7 (8.6)	1.14	0.19-6.63	0.888
>4.5 mm	99 (42.7)	9 (9.1)	1.20	0.21-6.81	0.837
Retention					0.409
Cemented	201 (72.0)	22 (10.9)	1		
Screwed	75 (26.9)	5 (6.7)	0.58	0.16-2.11	0.409
Overdenture	3 (1.1)	0	n/a	n/a	n/a
Splinted					
No	204 (72.9)	14 (6.9)	1		
Yes	76 (27.1)	13 (17.1)	2.80	0.98-8.02	0.055
					(Continues)

	Total, mean $\pm$ SD	Peri-implantitis,			
Characteristic	or n (%)	n (%)	OR	95% CI	<i>p</i> value
Number of annual maintenance visits during radiograph period (T2–T3)					0.079
≤1	63 (23.1)	5 (7.9)	1		
>1-≤2	104 (38.1)	4 (3.8)	0.46	0.11–1.96	0.296
>2-≤3	77 (28.2)	12 (15.6)	2.14	0.56-8.22	0.267
>3	29 (10.6)	5 (17.2)	2.42	0.44–13.2	0.309
Number of annual maintenance visits (T0-T4)					0.280
≤0.5	61 (22.4)	5 (8.2)	1		
>0.5-≤1	59 (21.7)	4 (6.8)	0.82	0.17-3.92	0.798
>1-≤1.5	91 (33.5)	6 (6.6)	0.79	0.16-3.95	0.775
>1.5	61 (22.4)	11 (18.0)	2.46	0.64–9.44	0.188

Note: p value by Wald test.

Abbreviations: CI, confidence interval; MTX, microtextured; OR, odds ratio; RBT, resorbable blast texturing; SLA, sandblasted, large-grit, acid-etched; T0, time of implant placement; T1, time of prosthetic restoration; T2, 1 year after prosthetic restoration; T3, time of exposure of last radiograph on which peri-implant bone could be clearly visualized; T4, time of last patient visit.

p < 0.05; p < 0.01; p < 0.01; p < 0.001.

**TABLE 3** Risk of incident peri-implantitis by patient, implant, and prosthesis characteristics during the radiograph period (T2–T3). Results from multivariate logistic regression with generalized estimation equations adjusting for duration and mean annual number of maintenance visits (N = 280 implants)

Characteristic	Total, mean $\pm$ SD or <i>n</i> (%)	Peri-implantitis, n (%)	OR	95% CI	p value
Number of implants	280	27 (9.6)			
Study group					
Non-exposed (0 threads exposed)	182 (65.0)	8 (4.4)	1		
Exposed (≥1 thread exposed)	98 (35.0)	19 (19.4)	7.82	1.91-32.0	0.004**
Patient age at T0, years	$63.0 \pm 11.3$		0.95	0.90-0.99	0.016*
Thread design					0.205
Buttress	46 (16.4)	2 (4.3)	1		
Progressive square	7 (2.5)	0	n/a	n/a	n/a
Reverse buttress	93 (33.2)	13 (14.0)	0.35	0.04–3.11	0.348
Square	20 (7.1)	5 (25.0)	2.02	0.26-15.9	0.506
V-shaped	114 (40.7)	7 (6.1)	0.23	0.20-2.28	0.211
Splinted					
No	204 (72.9)	14 (6.9)	1		
Yes	76 (27.1)	13 (17.1)	3.49	1.02–12.0	0.047*
Duration of radiograph period (T2-T3), years	$4.60 \pm 2.52$		1.19	0.95–1.50	0.136
Number of annual maintenance visits during radiograph period (T2–T3)					0.052
≤1	63 (23.1)	5 (7.9)	1		
>1-≤2	104 (38.1)	4 (3.8)	0.84	0.20-3.52	0.811
>2-≤3	77 (28.2)	12 (15.6)	3.23	0.57-13.9	0.114
>3	29 (10.6)	5 (17.2)	5.16	0.73-36.4	0.101

Note: p values by Wald test.

Abbreviations: CI, confidence interval; OR, odds ratio; T2, 1 year after prosthetic restoration; T3, time of exposure of last radiograph on which peri-implant bone could be clearly visualized.

p < 0.05; p < 0.01.

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only eight (2.9%) implants from this series failed. The overall implant level PI rate was 9.6% (and only 4.4% of the implants that did not have any interproximal threads exposed after the initial physiologic bone remodeling), which is well within, actually at the lower end of, the reported range between 0.4% and 85%.<sup>5,7,40,45-47</sup> Importantly, almost one-fifth (19.4%) of the implants with such thread exposure developed PI. This is the same overall rate as that found for implants placed by general practitioners.<sup>48</sup>

Our stringent eligibility criteria were selected to create the test and comparison groups for comparisons as precise and valid as possible. It requires a large source population to conduct such a study, which can be deducted from including only 165 patients from a pool of 4325 active patients whose charts were screened. The low eligibility rate of 3.8% also leads to potential bias in representing any real-life population. Hence, this study could be perceived as a proof-of-concept study, although the prevalence of PI corresponds to findings from non-academic studies. The paucity of such large, well-documented source populations may be a reason for the lack of studies like this. A main limitation of this study is the high number and great diversity in skill levels of various categories of providers as well as the variety of implant systems used, some of which have been associated with the prevalence of PI.<sup>26</sup> The same applies to the various prosthetic designs included, some of which may be considered risk indicators for PI.49

Furthermore, with this study being primarily based on radiographic assessment, the observed correlation between implant threads not embedded in bone and an increased risk for the onset of PI could not consider soft tissue variables, such as keratinized mucosa width, mucosal thickness, or peri-implant soft tissue height. Moreover, we could not assess the presence/absence of buccal thread exposure due to the utilization of two-dimensional radiographs allowing only assessment of the interproximal aspects. Finally, inherent in the study design are the limitations of any retrospective study, such as no new data being collected and the data having been recorded for purposes other than this study with no possibility for randomization and recording of prospective observations.

#### 5 CONCLUSION

Within the limitations of this retrospective study, and age being the only non-modifiable risk factor identified, splinting and implant thread exposure (no BIC) after the expected initial bone remodeling were the only statistically significant potentially modifiable risk indicators for incident PI that were identified in this study. Implants with  $\geq 1$ thread exposed 1 year after implant restoration were 7.82 times more likely to develop PI than those with no exposed threads. This impact occurred in a dose-response manner, as the risk for PI increased with increasing number of exposed threads, with each additional exposed thread increasing the risk of PI almost four-fold.

#### AUTHOR CONTRIBUTIONS

Study conception and design: Andrea Ravidà, Hom-Lay Wang, and Pablo Galindo-Moreno. Data collection: Andrea Ravidà, Ankita Samal, Musa Qazi, and Liana Preto Webber. Analysis and interpretation of the data: Hom-Lay Wang, Pablo Galindo-Moreno, Wenche S. Borgnakke, and Muhammad H. A. Saleh. Drafting of the manuscript: Andrea Ravidà, Liana Preto Webber, Wenche S. Borgnakke, and Muhammad H. A. Saleh. All authors gave their final approval and agreed to be accountable for all aspects of the work.

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#### CONFLICT OF INTEREST

All authors declare that they have no conflicts of interest related to this manuscript.

#### DATA AVAILABILITY STATEMENT

The data that support the findings of this study are available from the corresponding author upon request.

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#### SUPPORTING INFORMATION

Additional supporting information can be found online in the Supporting Information section at the end of this article.

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