# Extension and Density of Root Fillings and Postoperative Apical Radiolucencies in the Veterans Affairs Dental Longitudinal Study

Yan Zhong, DMD, PhD,\* Joel Chasen, DMD,<sup>†</sup> Ryan Yamanaka, DDS,<sup>†</sup> Raul Garcia, DMD, MMSc,<sup>†</sup> Elizabeth Krall Kaye, PhD, MPH,<sup>†</sup> Jay S. Kaufman, PhD,<sup>‡</sup> Jianwen Cai, PhD,<sup>‡</sup> Tim Wilcosky, PhD,<sup>‡</sup> Martin Trope, DMD,\* and Daniel J. Caplan, DDS, PhD<sup>f</sup>

#### **Abstract**

We evaluated the association between radiographically assessed extension and density of root canal fillings and postoperative apical radiolucencies (ARs) by using data from 288 participants in the Veterans Affairs Dental Longitudinal Study. Study subjects were not Veterans Affairs patients; all received their medical and dental care in the private sector. Generalized estimating equations were used to account for multiple teeth within subjects and to control for covariates of interest. Defective root filling density was associated with increased odds of postoperative AR among teeth with no preoperative AR (odds ratio, 3.0; 95% confidence interval [CI], 1.3-7.1), although preoperative AR was the strongest risk factor for postoperative AR (odds ratio, 29.2; 95% CI, 13.6-63.0 among teeth with ideal density). Compared with well-extended root fillings, neither overextended nor underextended root fillings separately were related to postoperative AR, but when those 2 categories were collapsed into one poorly extended category, poor extension was related to postoperative AR (odds ratio, 1.8; 95% CI, 1.1-3.2). (J Endod 2008;34: 798 – 803)

#### **Kev Words**

Endodontics, epidemiology, longitudinal studies, periapical periodontitis, root canal therapy

From the \*School of Dentistry and \*School of Public Health, University of North Carolina, Chapel Hill, North Carolina; \*Boston University Goldman School of Dental Medicine, Boston, Massachusetts; and \*College of Dentistry, University of Iowa, Iowa City, Iowa.

Reprints will not be available from the authors.

Address correspondence to Dr. Yan Zhong, 873 Hercules Lane, Foster City, CA 94404. E-mail address: yanzhong06@gmail.com.

0099-2399/\$0 - see front matter

Copyright © 2008 American Association of Endodontists. doi:10.1016/j.joen.2008.03.022

With regard to published studies of root canal therapy (RCT) and apical periodontitis (AP), several consistencies have emerged, including: AP and RCT are common (1.4%-8.5% and 1.5%-21.5% of teeth, respectively); AP is more prevalent among root canal filled (RCF) teeth than non-RCF teeth (16.7%-61.0%) of RCF teeth versus 0.53%-4.4% of non-RCF teeth); and overall quality of root fillings generally is poor (inadequate quality in 51%-86% of RCF teeth) (1-3). Criteria used to assess quality of root fillings often are based on the radiographically assessed characteristics of density (the extent to which the root filling material uniformly and completely fills the canals) and extension (the distance from the end of the root filling material to the radiographic apex). Root fillings designated as inadequate by these criteria do not necessarily lead to unsatisfactory endodontic results, and root fillings designated as adequate by these criteria do not necessarily lead to satisfactory endodontic results. Still, many follow-up clinical studies on RCF teeth suggest that poor root filling density and extension are associated with postoperative AP (4-7).

It is difficult to make direct comparisons between relevant articles because studies differ in many ways. Gaps in the literature include the following:

One gap relates to methodologic flaws. Correlations among teeth within individuals often are ignored (7–9). Analyses that do not account for clustering of observations (ie, teeth) within individuals ignore the appropriate correlation structure and can lead to incorrect inferences in hypothesis testing (10, 11), whereas analyses that randomly select one observation per person are inefficient because not all data are used.

Another gap relates to the limited populations studied. Conclusions made from epidemiologic studies generally are considered more valid if similar findings are observed among different populations and across different conditions. In one recent review (12), only one U.S. study (13) was included among the 11 identified observational cohort studies. In addition, all previous U.S. studies involved either patients from dental clinics or teaching hospitals (4, 13–16) or enrollees in dental insurance programs (17); none has used a population-based sample; thus, endodontic outcomes in the general U.S. population remain unstudied.

The third gap is that few longitudinal data are available. Compared with cariology and periodontology, epidemiologic data on endodontology are scarce. Cross-sectional designs or longitudinal designs with only 6–12 months of observation prevail among existing studies. Healing and growth of periapical lesions are dynamic processes that require considerable time; 3–4 years or more might be required to record a stable treatment outcome (18, 19).

To address these gaps we conducted the present study, the aim of which was to investigate the association between radiographically assessed extension and density of root fillings and the prevalence of apical radiolucency (AR) 3 years after RCT, while controlling for the presence of preoperative AR and other tooth-level and person-level covariates.

## **Material and Methods**

Data analyzed in this study were from the Veterans Affairs Dental Longitudinal Study (VADLS), an ongoing, closed-panel longitudinal study of oral health and disease

among 1,231 adult men aged 25–85 years at baseline. The cohort was established starting in 1968 through community-based recruitment of adult men from the greater Boston metropolitan area. Subjects were not patients of the VA system; rather, they received dental and medical care through the private sector.

One goal of the VADLS has been to identify determinants of oral health in an aging population. Participants had varying oral conditions at baseline, although all were free of chronic medical conditions. Since baseline, study participants have been seen once every 3 years for comprehensive dental and medical examinations. Dental examinations include both clinical and radiographic components. The clinical component records decayed, missing, or filled coronal tooth surfaces (DMFS) and periodontal status. The radiographic component includes a full mouth series of intraoral radiographs (20).

To date, the cohort has been under observation for more than 30 years. The average interval between VADLS exams has been approximately 38 months (21). The present study used the existing computerized VADLS database to identify a random sample of 853 dentate participants, each with a complete record for each of the 32 permanent teeth/tooth spaces at each examination cycle. The protocol for this study was reviewed and approved by the Institutional Review Board on Research Involving Human Subjects at the VA Boston Healthcare System.

Our analysis includes only incident RCT, ie, those teeth that had RCT initiated and completed after the baseline examination. For these teeth, follow-up started at the cycle when RCT was first detected from radiographs (ie, the index cycle) and ended at the next examination cycle. At the next cycle, if there was a radiolucency detected around the root apex, the outcome was classified as presence of postoperative AR, regardless of whether a lesion was present preoperatively or whether it developed after treatment.

Determinations of root filling extension and density, in addition to AR, were made solely from available radiographs of diagnostic quality. Two second-year endodontic residents from Boston University independently reviewed study subjects' intraoral radiographs. Before data collection, a training and calibration session for the radiographic examiners was conducted to assure adequate reliability of radiographic examination and to evaluate diagnostic criteria for endodontic assessments. Kappa values describing interexaminer reliability were excellent for the endodontic variables, ranging from 0.80-1.00, depending on the variable (22).

Diagnostic criteria for radiographic evaluation of AR, extension, and density were adapted from Odesjo et al. (23):

#### **AR (Both Preoperative and Postoperative)**

- Present: periapical rarefaction contiguous with periodontal ligament space >1 mm wide, with absence of intact lamina dura.
- Not present: apical periodontal ligament space <1 mm thick.

Multirooted teeth were classified as AR present if at least 1 root met the criteria for AR.

#### **Extension**

- Underextended: >2 mm short of the radiographic apex.
- Well-extended: 0-2 mm short of the radiographic apex.
- Overextended: root filling material beyond the radiographic apex.

Multirooted teeth were classified as well-extended only if all roots were well-extended, overextended if any roots were overextended, and underextended otherwise.

#### **Density**

- Ideal: no voids or inhomogeneous zones visible, no space discernible between filling material and canal wall.
- Acceptable: no voids or inhomogeneous zones visible in apical third, no space discernible between filling material and canal wall in apical third
- Defective: voids or inhomogeneous zone visible in apical third, or space is discernible between filling material and canal wall in apical third (also includes any teeth with unfilled canals).

Multirooted teeth were classified according to the root filling with the worst density.

In addition to data collected from intraoral radiographs, other variables were available from the parent VADLS data set and included in this study. On the basis of their temporal relationships, factors analyzed with respect to endodontic prognosis can be classified as preoperative (eg, presence of preoperative AR, patient age, income, education, smoking history, diabetes, hypertension, body mass index, tooth type), intraoperative (eg, extension and density of root filling, treatment-related complications, year of RCT, type of root filling material, number of posts, post orientation), and postoperative (eg, open access, crown) (18). Values for the covariates were taken from the index cycle for each tooth.

Statistical analyses were performed in 3 stages: (1) univariate description of data by using frequencies and percentages; (2) bivariate associations between the main exposure variables and the outcome, with preliminary assessment of potential interactions and confounders by using contingency tables and  $\chi^2$  tests of proportions; and (3) multivariable logistic regression analyses to evaluate associations in the presence of other factors. Postoperative AR was the dichotomous dependent variable, with extension and density as explanatory variables of primary interest. Preoperative AR was analyzed as an effect modifier to allow calculation of separate estimates of effect depending on whether a tooth had preoperative AR, whereas other covariates were analyzed as potential confounders. Generalized estimating equations (GEE) were used to adjust for correlation of teeth within subjects (24, 25). All statistical tests were two-tailed and performed with SAS Version 9.1 (SAS Institute Inc, Cary, NC).

#### Results

Eight hundred fifty-three participants contributed a total of 27,296 teeth or tooth spaces (Fig. 1). The present analysis included only the 609 teeth that received RCT after baseline and had complete data at 3 consecutive cycles (ie, the cycle preceding the index cycle; the index cycle; and the very next cycle). Two hundred eighty-eight unique individuals contributed to the final sample, with the number of teeth per subject ranging from 1–11 (Table 1).

Of the 609 analyzed teeth, 68 (11.2%) had postoperative AR at the end of follow-up (Table 2). Ideal density and adequate extension were observed in 216 teeth (35.5%), leaving 393 (64.5%) with unsatisfactory root filling quality. Preoperative AR existed in 79 teeth (13.0%). The mean age of participants at the time of RCT for each tooth was 61 years. RCT-related complications were noted infrequently: perforation was seen in only 0.5%, broken instruments in 1.2%, unfilled canals in 1.3%, unfilled roots in 1.6%, and insufficient length of root filling (<4 mm of root filling material in the most apical portion of the root canal) in 2.1% of teeth, respectively. In addition, only 3 teeth had open access, and 11 teeth were from individuals who had diabetes. Because these occurrences were few, these variables were not analyzed further.

Bivariate analyses (data not shown) revealed defective density, preoperative AR, and off-axis post orientation to be significantly asso-

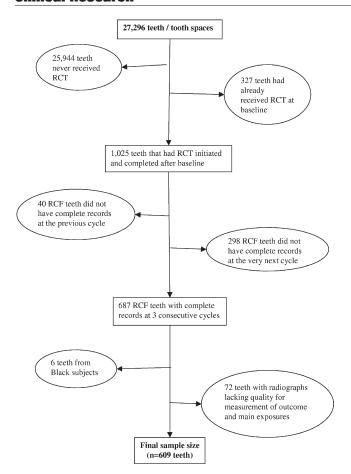


Figure 1. Description of study sample.

ciated with postoperative AR (p < .05). Stratified analyses suggested an interaction between defective density and preoperative AR, and preliminary assessment of potential confounders indicated that filling material, post orientation, the presence of a crown, tooth type, age, income, and smoking history were potential confounders.

In the final model (Table 3), defective density remained significantly associated with increased odds of postoperative AR after controlling for extension and preoperative AR (p = .028). The significant interaction term between defective density and preoperative AR (p < .0001) demonstrated that the effect of defective density on the odds of postoperative AR differed depending on whether the tooth had preoperative AR. When using teeth with ideal density and no preoperative AR

**TABLE 1.** Number of RCF Teeth Contributed by Subjects

No. of RCF Teeth Contributed Per Subject	No. of Subjects (%)	Total No. of RCF Teeth Contributed
1	153 (53)	153
2	58 (20)	116
3	32 (11)	96
4	18 (6)	72
5	9 (3)	45
6	10 (4)	60
7	3 (1)	21
8	1 (0)	8
9	3 (1)	27
10	0 (0)	0
11	1 (0)	11
Total	288 (100)	609

**TABLE 2.** Univariate Distribution of Study Variables (N = 609 Teeth)

Variables	Level	Frequency	Percent
Postoperative AR	Yes	68	11.2
(outcome)	No	541	88.8
Extension (main	Underextended	85	14.0
exposure)	Overextended	122	20.0
•	Well-extended	402	66.0
Density (main	Acceptable	154	25.3
exposure)	Defective	158	25.9
·	Ideal	297	48.8
Preoperative AR	Yes	79	13.0
	No	530	87.0
Filling material	Gutta-percha only	556	91.3
•	Silverpoint or both	53	8.7
No. of posts	≥1	318	52.2
•	0	291	47.8
Post orientation	Off axis	17	2.8
	On axis	592	97.2
Crown	Yes	344	56.5
	No	265	43.5
Tooth type	Premolar	213	35.0
	Molar	184	30.2
	Anterior	212	34.8
Income*	>\$25,000 per year	200	32.8
	≤\$25,000 per year	397	65.2
Education	College graduate	191	31.4
	Some college	252	41.4
	High school or less	166	27.3
Smoking history	Current smoker	231	37.9
	Former smoker	201	33.0
	Never smoker	177	29.1
Hypertension	Yes	79	13.0
	No	530	87.0
Year of RCT	≥1985	270	44.3
	<1985	339	55.7
Age at access	≥65	213	35.0
(years)	55–64.9	245	40.2
	<55	151	24.8
Body mass index*	≥25 kg/m <sup>2</sup>	369	60.6
	13-24.9 kg/m <sup>2</sup>	239	39.2

<sup>\*</sup>n does not add up to 609 because of missing values.

as the reference group, odds ratios (95% confidence interval [CI]) of postoperative AR were 3.0 (1.3–7.1) for teeth with defective density but no preoperative AR, 29.2 (13.6-63.0) for teeth with ideal density and preoperative AR, and 33.2 (12.0-92.0) for teeth with both defective density and preoperative AR (Table 4). Controlling for the significant interaction term, neither acceptable density (relative to ideal density) nor extension was significantly associated with postoperative AR. However, when the underextended and overextended groups were combined into a single poorly extended category, a significant association between poor extension and odds of postoperative AR was observed, when compared with well-extended root fillings (odds ratio, 1.8; 95%) CI, 1.1–3.2). Variables under investigation but not retained in the final model as a result of nonsignificance and minimal confounding included type of root filling material, post orientation, presence of crown, tooth type, age at access, income, smoking history, number of posts, year of RCT, education, hypertension, and body mass index.

Finally, because the impact of preoperative AR was so strong, we performed an analysis restricted to the 530 teeth without preoperative AR (Table 5). In this model, defective density remained the only explanatory variable significantly associated with increased odds of postoperative AR (p=.022). Here, the combined category of poor extension conferred no significantly increased odds of postoperative AR (odds ratio, 1.8; 95% CI, 0.8–3.7). Income remained in the model as a result of its confounding effect on the main effects of interest, but its large p value (.798) indicates that income itself was not significantly predictive of postoperative AR.

**TABLE 3.** Final Multivariable Logistic Regression Model of Associations between Explanatory Variables and Postoperative AR (N = 609 Teeth)

Independent Variable	Level	p Value	Odds Ratio (95% CI)
Extension	Underextended	.199	1.7 (0.8–3.4)
	Overextended	.082	2.0 (1.0-3.9)
	Well-extended (reference)	_	· <u> </u>
Density	Acceptable	.290	1.5 (0.7–3.2)
•	Defective	.028*	See Table 4
	Ideal (reference)	_	_
Preoperative AR	Yes	<.0001*	See Table 4
·	No (reference)	_	_
Defective density, preoperative AR	See Table 4	<.0001*	See Table 4

<sup>\*</sup>Statistically significant at .05 level.

# **Discussion**

After controlling for preoperative AR, defective density remained independently, significantly associated with postoperative AR. Voids in the root filling represent spaces that residual microflora could inhabit and subsequently transport endotoxins to the root apex, stimulating an inflammatory response (4). Our finding of significant interaction between preoperative AR and defective density suggests that the effect of defective density differs depending on whether preoperative AR exists; although our results confirm the importance of homogeneously dense root fillings, this especially is important when the tooth does not exhibit preoperative AR.

Underextended or overextended root fillings might lead to a poorer prognosis compared with well-extended root fillings, but our analyses failed to show significant associations between either underextension or overextension and postoperative AR when other factors were considered simultaneously, no matter whether preoperative AR existed or not. This finding agrees with some previous studies (26, 27) but differs from others (8, 14), and there are several possible explanations. In the case of underextension, if space is left apical to the tip of the filling material but the apex is free of bacteria, underextension is unlikely to increase the risk of periapical inflammation. In the case of overextension, a small amount of excess root filling material extruded into the periapical area might not be enough to induce a significant foreign body reaction. In addition, the number of teeth in the underextended or overextended group in our sample might be too small for statistically significant influences to be detected. To address this hypothesis, we collapsed underextended and overextended teeth into a single category and found that poor extension conferred significantly increased odds of postoperative AR compared with teeth with well-extended root fillings, but only in the presence of preoperative AR; this would be consistent with inadequate eradication of bacteria from the root canal space. Radiographic assessment of root filling extension and density is more easily quantified than antibacterial management, which might have resulted in an overemphasis on these factors in some studies.

In the final regression model, preoperative AR was of greater significance than other variables evaluated. This makes intuitive sense because (1) teeth with preoperative AR presumably already have bacteria present in the apical regions of their root canal systems, whereas teeth without preoperative AR might or might not have bacteria present

**TABLE 4.** Assessment of Interaction between Defective Density and Preoperative AR in the Final Model (N = 609 Teeth)

Odds Ratio For Postoperative AR (95% CI)		Preoperative AR		
		Yes	No	
Density	Defective Ideal	33.2 (12.0–92.0) 29.2 (13.6–63.0)	3.0 (1.3–7.1) 1.0 (reference)	

there; and (2) eradicating bacteria already present throughout the root canal system is more difficult than preventing bacteria from ever reaching the apical regions of the root. However, our study found a much greater impact of preoperative AR on prognosis than that described previously. This could be explained in part by differences in diagnostic criteria used to define AR between the present study and other studies (28-30) or by differences in the study population and study design (31). For example, the present study participants are relatively older, so their ability to repair preexisting periapical lesions might be less than in younger populations, and thus they might be at greater risk for radiographically evident postoperative inflammation (32). It might take longer for an older host to heal an existing lesion, even after wellperformed RCT; it has been reported that delayed healing of periapical radiolucency could occur as long as 27 years after treatment (19). The 3-year follow-up in our study might not have been sufficient for some preoperative ARs to heal completely.

In the presence of preoperative AR, the effect of other factors investigated here seems less important, consistent with other studies (29, 33, 34). There are several possible explanations for this consistency. First, many investigated factors truly might not have a confounding effect. Second, the study might have insufficient power to detect relatively small effects associated with certain covariates. Finally, some covariates were excluded from regression modeling procedures because they either had too imbalanced a distribution or too few occurrences to provide meaningful point estimates and CIs. Regardless, for most factors under investigation, the 288 subjects and 609 teeth provided a large enough sample for GEE methodology to produce consistent estimates (25).

A total of 327 RCF teeth were excluded from the analysis because they already had received RCT before baseline (Fig. 1). Additional analyses (not shown here) revealed no meaningful differences between these 327 teeth and the 1025 teeth that received RCT after baseline, except that the excluded teeth tended to be from older subjects and were

**TABLE 5.** Final Multivariable Logistic Regression Model of Associations between Explanatory Variables and Postoperative AR (Restricted to N=530 Teeth without Preoperative AR)

Independent Variable	Level	p Value	Odds Ratio (95% CI)
Extension	Underextended	.460	1.5 (0.6–3.7)
	Overextended	.197	2.0 (0.8-5.1)
	Well-extended (reference)	_	_
Density	Acceptable	.467	1.5 (0.5-4.6)
•	Defective	.022*	3.2 (1.3-8.2)
	Ideal (reference)	_	_
Income	>\$25,000	.798	0.9 (0.4-2.0)
	≤\$25,000 (reference)	_	

<sup>\*</sup>Statistically significant at .05 level.

more likely to be anterior teeth. The prevalence of postoperative AR (11.2%) among the 609 analyzed RCF teeth is slightly lower than the frequencies of AR reported in the literature review. This could be partly because our study population is a community-based cohort, whereas previous studies mostly involved clinic patients. The satisfactory quality of root filling in this sample (35.5%) is comparable with the reported rate in the literature (ie, 30%-40%), which emphasizes the need for improving root filling quality in endodontic practice.

Only white men were included in the present study sample, so caution should be exercised when generalizing these results to other populations. However, because subjects received dental and medical care in the private sector, this sample likely was more socially diverse than participants recruited in many clinical studies, such as those conducted in dental schools or through dental insurance programs (4, 13–17). It is noteworthy, however, that there have been no consistent findings in the literature to indicate that root filling quality or outcome varies by race, ethnicity, or sex.

As with all retrospective studies, data quality was dependent on the availability and completeness of documentation. Data collection for many variables was restricted to available information from the electronic database, and certain variables that might be related to postoperative AR were not available (eg, bacterial levels in the root canal space and adequacy of the coronal restoration). With respect to the collected endodontic variables, potential misclassifications exist for several reasons. First, endodontic variables were assessed on the basis of radiographs, and the use of 2-dimensional radiographs to represent 3-dimensional structures can be problematic in quantifying variables such as root filling extension and density. Second, this study operated under the assumption that postoperative AR is an undesired health outcome, given the general correlation between AR and histologically confirmed inflammatory status of the periapical tissues (35). Third, assessment of periapical health did not involve clinical evaluation of study subjects. Although postoperative ARs are not always associated with clinical symptoms (12), not all ARs represent chronic AP, they could instead be periapical cysts, foreign body reactions, or scar tissue. Finally, radiographic assessment is an uncertain process with wide variations among dentists, but this concern was minimized in our study by employing similarly trained examiners and calibrating them before data collection.

In conclusion, the major findings from this follow-up study were that: (1) defective root filling density was associated with increased odds of postoperative AR, but only among teeth without preoperative AR; (2) the most significant risk factor for postoperative AR was the presence of preoperative AR; (3) the statistical significance of root filling extension was dependent on how the variable was classified; and (4) none of the other tooth-level or person-level covariates assessed was significantly related to postoperative AR. Failure of endodontic treatment is caused by microbes and their by-products being introduced into or remaining in the root canal system and periapical region, and root fillings with inadequate extension or density can only facilitate the infectious process.

# **Acknowledgments**

The authors thank Drs Jeffrey Hutter and John Stamm, without whose assistance this study would not have been possible. The VA Dental Longitudinal Study and the VA Normative Aging Study are components of the Massachusetts Veterans Epidemiology Research & Information Center, VA Boston Healthcare System, Boston, MA. The studies are supported by the VA Cooperative Studies Program/ERIC, U.S. Department of Veterans Affairs. Dr Garcia was the recipient of a Career Development Award in Health Services Research from VA HSR&D Service. He is supported by a VA Epidemiology Merit

Review Award and by NIH grant K24-DE00419 from the National Institute of Dental and Craniofacial Research. The study also was supported by NIDCR Grant R01-DE13807.

### References

- Soikkonen KT. Endodontically treated teeth and periapical findings in the elderly. Int Endod J 1995;28:200 –3.
- Weiger R, Hitzler S, Hermle G, Lost C. Periapical status, quality of root canal fillings and estimated endodontic treatment needs in an urban German population. Endod Dent Traumatol 1997;13:69–74.
- Marques MD, Moreira B, Eriksen HM. Prevalence of apical periodontitis and results of endodontic treatment in an adult, Portuguese population. Int Endod J 1998;31:161–5.
- Ray HA, Trope M. Periapical status of endodontically treated teeth in relation to the technical quality of the root filling and the coronal restoration. Int Endod J 1995;28:12–8.
- Kirkevang LL, Orstavik D, Horsted-Bindslev P, Wenzel A. Periapical status and quality
  of root fillings and coronal restorations in a Danish population. Int Endod J
  2000;33:509–15
- Tronstad L, Asbjornsen K, Doving L, Pedersen I, Eriksen HM. Influence of coronal restorations on the periapical health of endodontically treated teeth. Endod Dent Traumatol 2000:16:218–21.
- Hommez GM, Coppens CR, De Moor RJ. Periapical health related to the quality of coronal restorations and root fillings. Int Endod J 2002;35:680–9.
- Sjogren U, Hagglund B, Sundqvist G, Wing K. Factors affecting the long-term results of endodontic treatment. J Endod 1990;16:498–504.
- Farzaneh M, Abitbol S, Lawrence HP, Friedman S. Treatment outcome in endodontics—the Toronto Study. Phase II: initial treatment. J Endod 2004;30: 302-9.
- Kleinbaum DG, Klein M. Logistic regression: a self-learning text. 2nd ed. New York: Springer-Verlag Inc, 2002.
- Caplan DJ, Slade GD, Gansky SA. Complex sampling: implications for data analysis. J Public Health Dent 1999;59:52–9.
- 12. Friedman S. Prognosis of initial endodontic therapy. Endod Topics 2002;2:59 88.
- 13. Trope M, Delano EO, Orstavik D. Endodontic treatment of teeth with apical periodontitis: single vs multivisit treatment. J Endod 1999;25:345–50.
- Buckley M, Spangberg LS. The prevalence and technical quality of endodontic treatment in an American subpopulation. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 1995;79:92–100.
- Chugal NM, Clive JM, Spangberg LS. A prognostic model for assessment of the outcome of endodontic treatment: effect of biologic and diagnostic variables. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2001;91:342–52.
- Chugal NM, Clive JM, Spangberg LS. Endodontic infection: some biologic and treatment factors associated with outcome. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2003;96:81–90.
- Salehrabi R, Rotstein I. Endodontic treatment outcomes in a large patient population in the USA: an epidemiological study. J Endod 2004;30:846-50.
- Friedman S. Treatment outcome and prognosis of endodontic therapy. In: Ørstavik D, Pitt Ford TR., eds. Essential endodontology: prevention and treatment of apical periodontitis. Oxford: Blackwell Science Ltd, 1998:367–401.
- Fristad I, Molven O, Halse A. Nonsurgically retreated root filled teeth: radiographic findings after 20–27 years. Int Endod J 2004;37:12–8.
- Garcia RI, Krall EA, Vokonas PS. Periodontal disease and mortality from all causes in the VA Dental Longitudinal Study. Ann Periodontol 1998;3:339 – 49.
- Krall EA, Garvey AJ, Garcia RI. Alveolar bone loss and tooth loss in male cigar and pipe smokers. J Am Dent Assoc 1999;130:57–64.
- Landis JR, Koch CG. The measurement of observer agreement for categorical data. Biometrics 1977:33:159-74.
- Odesjo B, Hellden L, Salonen L, Langeland K. Prevalence of previous endodontic treatment, technical standard and occurrence of periapical lesions in a randomly selected adult, general population. Endod Dent Traumatol 1990;6:265–72.
- Liang KY, Zeger SL. Longitudinal data analysis using generalized linear models. Biometrika 1986;73:13–22.
- Stokes ME, Davis CS, Koch GG. Categorical data analysis using the SAS system. 2nd ed. Cary, NC: SAS Institute Inc, 2000.
- Bystrom A, Happonen RP, Sjogren U, Sundqvist G. Healing of periapical lesions of pulpless teeth after endodontic treatment with controlled asepsis. Endod Dent Traumatol 1987;3:58–63.
- Sjogren U, Figdor D, Persson S, Sundqvist G. Influence of infection at the time of root filling on the outcome of endodontic treatment of teeth with apical periodontitis. Int Endod J 1997;30:297–306.

# **Clinical Research**

- Ørstavik D, Qvist V, Stoltze K. A multivariate analysis of the outcome of endodontic treatment. Eur J Oral Sci 2004;112:224–30.
- 29. Friedman S, Abitbol S, Lawrence HP. Treatment outcome in endodontics: the Toronto Study. Phase 1: initial treatment. J Endod 2003;29:787–93.
- Dugas NN, Lawrence HP, Teplitsky PE, Pharoah MJ, Friedman S. Periapical health and treatment quality assessment of root-filled teeth in two Canadian populations. Int Endod J 2003;36:181–92.
- Caplan DJ. Epidemiologic issues in studies of association between apical periodontitis and systemic health. Endodontic Topics 2004;8:15–35.
- Ashcroft GS, Mills SJ, Ashworth JJ. Ageing and wound healing. Biogerontology 2002;3:337–45.
- Aleksejuniene J, Eriksen HM, Sidaravicius B, Haapasalo M. Apical periodontitis and related factors in an adult Lithuanian population. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2000;90:95–101.
- Kirkevang II., Wenzel A. Risk indicators for apical periodontitis. Community Dent Oral Epidemiol 2003;31:59-67.
- Ørstavik D, Kerekes K, Eriksen HM. The periapical index: a scoring system for radiographic assessment of apical periodontitis. Endod Dent Traumatol 1986;2:20–34.