

Morphologic Variations and Gender Dimorphism in Mandibular First Premolar Root Canal Pattern: A CBCT Study in South Indian Population

Nidhi James^{1-a}, Tina Puthen Purayil^{1-b*}, Vathsala Patil^{2-c}, Kalyana Chakravarthy Pentapati^{3-d}, Ravindranath Vineetha^{2-e}

¹ Department of Conservative Dentistry and Endodontics, Manipal College of Dental Sciences, Manipal, Manipal Academy of Higher Education, Manipal, Karnataka, India

² Department of Oral Medicine and Radiology, Manipal College of Dental Sciences, Manipal, Manipal Academy of Higher Education, Manipal, Karnataka, India
³ Department of Public health dentistry, Manipal College of Dental Sciences, Manipal, Manipal Academy of Higher Education, Manipal, Karnataka, India

^{*}Corresponding author

| Research Article | ABSTRACT | | | | | | | | | | |
|----------------------------------|--|--|---|--|--|--|--|--|--|--|--|
| History | | | f mandibular first premolars (MFP) in south | | | | | | | | |
| HISTORY | 0 | Indians and also evaluated the gender dimorphism and contralateral symmetry of pattern. Methods : 200 Cone Beam Computed Tomography (CBCT) images of MFP of patients above 18 years were | | | | | | | | | |
| Received: 26/06/2022 | analysed for the root canal morphology after excluding the distorted images, endodontically treated tooth or | | | | | | | | | | |
| Accepted: 02/02/2023 | fractured root. The teeth were carefully evaluated in cross sectional, axial and coronal sections and the | | | | | | | | | | |
| | | | oot canal as per Vertucci's classification. | | | | | | | | |
| | Interobserver and intraobserver relia | | • | | | | | | | | |
| | Results: Single rooted teeth with Typ | e I (73.01%) variation was the r | most prevalent pattern observed in the | | | | | | | | |
| License | study population, which was followed | d by Type III (14.11%) and Type | e V (5.52%), respectively. Type IV and Type | | | | | | | | |
| | VIII variations were not found in our | , | | | | | | | | | |
| | | | e most common pattern with symmetry in | | | | | | | | |
| This work is licensed under | contralateral mandibular single-roote | ed first premolars. | | | | | | | | | |
| Creative Commons Attribution 4.0 | | | | | | | | | | | |
| International License | Key words: Cone Beam Computed To | mography, Mandible, Morpho | logy, Premolar, Root Canal. | | | | | | | | |
| | | | | | | | | | | | |
| 2 mary.james.nj@gmail.com | Dhttps://orcid.org/0000-0002-5351-5150 | ▶ 🔄 tina.pp@manipal.edu | Dhttps://orcid.org/0000-0002-8887-2297 | | | | | | | | |
| | Dhttps://orcid.org/0000-0002-8656-8080 | d 😆 kalyan.cp@manipal.edu | Dhttps://orcid.org/0000-0002-5462-5677 | | | | | | | | |
| evineetha.manu@manipal.edu | Dhttps://orcid.org/0000-0002-8932-9982 | | | | | | | | | | |

How to Cite: James N, Purayil TP, Patil V, Pentapati KC, Vineetha R. (2023) Morphologic Variations and Gender Dimorphism in Mandibular First Premolar Root Canal Pattern: A CBCT Study in South Indian Population, Cumhuriyet Dental Journal, 26(1):17-21.

Introduction

The maxillary and mandibular premolar teeth are known to have significant morphological variations in root canals compared to other teeth. It is reported more commonly in maxillary second premolars and mandibular first premolars.¹⁻⁴ A thorough knowledge and understanding of root canal morphology(RCM) is vital for the success of root canal therapy.^{3,5,6}

Mandibular first premolars (MFP) have two pulp horns; a large buccal horn with sharp outline and an inconspicuous round lingual horn.⁷ Access to the buccal root canal is easily achieved, whereas the lingual canal access is complicated due to the deviating path. The lingual inclination of these teeth also makes it difficult to locate the lingual orifice as the file insertion will be directed towards the buccal aspect of the tooth⁸. Moreover, frequent morphologic variations like C-shaped canals and extra root canals cause challenges in the biomechanical preparation and in obturation phases.^{9–15}

Various methods have been used to identify and record the root canal morphologies, some of which are two-dimensional radiographs, staining methods for root canals, sectioning of the dental hard tissues, micro-CT, and CBCT (Cone-Beam Computed Tomography) scanning. CBCT is superior to traditional radiographs and other digital imaging methods.¹⁶ The images procured by the CBCT are depicted in axial, sagittal, and coronal sections, and it also shows reduced superimposition by adjacent structures.

There is a general lack of literature on the variations in RCM of MFP in the south Indian population.^{9,17–20} Hence, we aimed to evaluate the variations in RCM of MFP in the south Indian subpopulation. We also aimed to compare the root canal configurations with gender and symmetry considering the number of roots in MFP. The study aimed to evaluate the variations in the RCM of mandibular first premolars (MFP) in south Indians and also evaluated the gender dimorphism and contralateral symmetry of pattern.

Material and Methods

A retrospective study on 200 CBCT images with MFP was conducted. We included CBCT images of individuals aged above 18 years with MFP. We have excluded images that were distorted, images with MFP which had endodontic treatment, fractured root, incomplete root formation, coronal or post endodontic restorations, and

physiological or pathological processes such as root resorption. A well-experienced radiologist performed CBCT imaging following the recommended protocol with the minimum exposure necessary for adequate image quality. CBCT images were obtained with i-CAT 17-19 Imaging System (Imaging Sciences International, USA). Exposure parameters were further adjusted according to patient indication for imaging. Image enhancement tools like zooming, contrast, and brightness adjustments, were used for better visualization and adjustment. Coronal, cross-sectional, and axial sections of the images were used to analyze the root morphology pattern. This study was conducted in the department of Oral Medicine and Radiology. The study protocol was approved by the institutional ethics committee, Kasturba Medical college and Kasturba Hospital, Manipal, India (Decision date-8.10.2019, IEC no- 726/2019)

The CBCT images were assessed using Anatomage software under standard illumination conditions by a single trained radiologist. Repeat assessment of 10% images was done after one month. Also, 10% of the images were randomly evaluated by an experienced radiologist. A Kappa value of 0.7 was obtained, showing substantial inter and intra-observer reliability. Each image was studied based on the number of roots and the canal morphology in each root according to the Vertucci's classification.²¹

All the analysis was done using SPSS version 20. A pvalue of <0.05 was considered statistically significant. Right, and left side variations, intra, and inter-examiner reliability were assessed using Kappa Coefficient. Comparison of RCM with sex was done using Fisher's exact test.

Results

A total of 200 MFP were assessed for their RCM, out of which 102 were left MFP, 166 had single root, 33 had two roots, and one had three roots (Table 1). In both right and left MFP, Vertucci's Type-I was the most common root canal configuration present (69.6% and 93% respectively), followed by and Type V (12.5%) on the right side and Type III (12.3%) on the left side (Table 1).

In single-rooted teeth, the right and left distribution of various RCM was compared using the Kappa coefficient. There was a 74% similarity on the right and left side distribution of RCM (Kappa=0.39). The right and left distribution of RCM was done independently for buccal and lingual root canals in two rooted teeth. In the buccal root, the similarity was 54.54% (Kappa=0.058). However, a high similarity was seen in the lingual root (81.81%; Kappa = 0.421).

Significant differences were seen in the distribution of RCM of single-rooted teeth on both the sides. In both right and left MFP, the distribution of the type I pattern was significantly higher in females than males (P=0.046 and 0.041), respectively (Table 2). However, no significant differences were seen in the distribution of RCM of two rooted teeth with sex on both the sides in buccal and lingual root canals (Table 3).

Discussion

The number of roots and root canal morphology in human population shows great diversity. Previous studies have utilized different methodologies to assess the root canal morphology and have reported complex internal anatomy of the MFP. Thorough knowledge of such variations in every population is indispensable. The advances in dental materials and diagnostic tools have along with thorough knowledge of canal morphology have increased the success rate of endodontic therapy, even in complex root canal configurations.^{17,22}

Literature shows the presence of extra root and extra canals in the mandibular first premolar.^{23,24} Present study showed that 79.8% of the teeth had a single root and 16.7% had two roots, and only one tooth had three roots. These observations were similar to previous studies. Studies done in India have reported that single RCM is the most common canal morphology.^{19,25} Sert and Bayirli reported 60.5% with a single canal and 39.5% with two or more canals.³ Singh and Pawar reported that 76% had a single canal, 22 had two canals, and 2% had three canals.²⁵ Trope *et al.* showed 89.1% to have single-rooted teeth, 10.9% to have two rooted teeth.²³ Yu *et al.* reported that 98% had a single root, and 2% had two roots.⁹ Yang *et al.* reported 77.14% had single-rooted canals.²⁶

Our study showed that Type I configuration (85.5%) is the most common pattern in MFP, which was in line with observations from the previous researches (50-88%) (Table 4).^{3,9,27–35,10,17–21,25,26} This was followed by Type III configuration (13%), which was similar to study by Sert and Bayirli among the Turkish population.³ However, these studies have reported that the 2nd most common pattern was C-shaped morphology. On the contrary present study, C-shaped configuration was noted in only 1% of the sample (Table 4).

Very few studies in the literature have evaluated the significant differences in RCM with respect to gender and quadrant wise distribution. We observed type I configuration to be the most commonly present configuration in both males and females on the right and left sides, which was in accordance to earlier reports.^{3,33} The symmetry of RCM in contralateral premolars has been studied previously.^{33,36,37} Our study also showed that in single-rooted teeth, there was 74% symmetry, while in two rooted teeth, it was only 55% in buccal and 82% in lingual root canals. This was lower than previous studies.^{33,37} While one study reported that there were only a few pairs that showed such symmetry.³⁶

Our sample showed one 3-rooted tooth, and 1% of the sample showed C-shaped canals. Previous studies have reported diverse and miscellaneous configurations like circumferential canals³⁰, Types like 1-3, 1-2-3, 2-1-3, and 2-1-2-1, ³¹ types 2-3, and 1-4.²⁰ However, such patterns were not reported in our sample. In both males and females, type I configuration was the most common pattern with symmetry in contralateral mandibular single-rooted first premolars. We used CBCT images to assess the variations in the RCM, which is superior to conventional

radiography and digital radiographic techniques. It is validated valuable tool to evaluate the complex canal morphology to improve the outcomes of endodontic therapy.

Conclusions

In the present study using CBCT, Vertucci's Type I root canal configuration was the most common variant followed by Type III configuration and there was no difference noted in canal morphology variations when evaluated quadrant wise. Further studies on larger and different sub-populations are required to understand the canal morphological variations for better endodontic treatment outcomes and good prognosis.

Acknowledgments

The authors would like to thank the Department of Oral Medicine and Radiology for their technical support in the study

Conflicts of Interest

There are no conflicts of interest.

Table 1. Sample characteristics, number of roots and root canal morphology

| | Number of | Numb | er of ro | oots | Root canal Morphology | | | | | | | |
|-------|-----------|------|----------|------|-----------------------|---------|-----------|----------|---------|----------|--------------|--|
| | teeth 1 | | | 3 | Туре І | Type II | Type III | Type V | Type VI | Type VII | C- shaped | |
| Left | 102 | 82 | 20 | 0 | 93 (76.2) | 2 (1.6) | 15 (12.3) | 8 (6.6) | 0(0) | 3(2.5) | 1(0.8) | |
| Right | 98 | 84 | 13 | 1 | 78(69.6) | 3(2.7) | 11(9.8) | 14(12.5) | 2(1.8) | 3(2.7) | 1(0.9) | |
| Total | 200 | 166 | 33 | 1 | 171 | 5 | 26 | 20 | 2 | 6 | 2 | |

Table 2. Comparison of root canal morphology with sex on right and left side in single rooted premolars (n=77)

| Se | P-value | |
|----------|--|--|
| Male | Female | r-value |
| N(%) | N(%) | |
| 27(69.2) | 34(89.5) | |
| 1(2.6) | 1(2.6) | |
| 8(20.5) | 1(2.6) | 0.041; Sig |
| 1(2.6) | 0(0) | 0.041, 31g |
| 2(5.1) | 1(2.6) | |
| 0(0) | 1(2.6) | |
| | | |
| 24(61.5) | 30(78.9) | |
| 2(5.1) | 1(2.6) | |
| 6(15.4) | 4(10.5) | |
| 4(10.3) | 0(0) | 0.046; Sig |
| 0(0) | 2(5.3) | |
| 3(7.7) | 0(0) | |
| 0(0) | 1(2.6) | |
| | Male N(%) 27(69.2) 1(2.6) 8(20.5) 1(2.6) 2(5.1) 0(0) 24(61.5) 2(5.1) 6(15.4) 4(10.3) 0(0) 3(7.7) | $\begin{array}{c c} N(\%) & N(\%) \\ 27(69.2) & 34(89.5) \\ 1(2.6) & 1(2.6) \\ 8(20.5) & 1(2.6) \\ 1(2.6) & 0(0) \\ 2(5.1) & 1(2.6) \\ 0(0) & 1(2.6) \\ \end{array}$ |

Table 3. Comparison of root canal morphology with sex on right and left side in double rooted premolars (n=11)

| | | S | P-value | | | |
|---------|----------|---------|---------|---------|--|--|
| | | Male | Female | r-value | | |
| Buccal | | N(%) | N(%) | | | |
| Left | Type I | 5(83.3) | 4(80) | >0.99 | | |
| Len | Type III | 1(16.7) | 1(20) | | | |
| Right | Туре І | 3(50) | 4(80) | 0.545 | | |
| Nigili | Type V | 3(50) | 1(20) | 0.343 | | |
| Lingual | | | | | | |
| | Туре І | 4(66.7) | 5(100) | | | |
| Left | Type III | 1(16.7) | 0(0) | - | | |
| | Type V | 1(16.7) | 0(0) | | | |
| Right | Type I | 4(66.7) | 5(100) | 0.455 | | |
| | Type V | 2(33.3) | 0(0) | 0.455 | | |

| Table 4. Review of root can | I morphology | from earlier studies |
|-----------------------------|--------------|----------------------|
|-----------------------------|--------------|----------------------|

| | | | | Method | Root canal morphology (%) | | | | | | | | | |
|------------------------|------|-----------------|----------|--------------------|---------------------------|------|------|------|-------|------|-----|------|---|-------------|
| Author | Year | Population | N | used | T | П | Ш | IV | V | VI | VII | VIII | Miscellaneous | C shaped |
| 1. Pineda et al | 1972 | Mexico | 20 2 | Radiograph | 69.3 | 0 | 4.9 | 1.5 | 23.4 | 0 | 0 | 0 | | |
| 2. Zillich & Dowson | 1973 | US | 139 3 | Radiograph | 66 | 5.2 | 0 | 17.5 | 0 | 0 | 0 | 0.4 | 7.6^ | |
| 3. Vertucci | 1984 | US | 400 | D&C | 70 | 0 | 4 | 1.5 | 24 | 0 | 0 | 0 | | |
| 4. Baisden et al | 1992 | US | 100 | Sectioning | 76 | 0 | 0 | 24 | 0 | 0 | 0 | 0 | | |
| 5. Caliskan et al | 1995 | Turkey | 100 | D&C | 64.15 | 7.55 | 3.77 | 7.55 | 9.43 | 1.89 | 0 | 5.66 | | |
| 6. Sert & Bayirli | 2004 | Turkey | 200 | D&C | 60.5 | 18.5 | 11.5 | 7 | 2.5 | 0 | 0 | 1 | | |
| 7. lyer et al* | 2006 | India | 200 0 | RVG | 75.4 | 1 | | 20.8 | 2.4 | 0 | 0 | 0.4 | | |
| 8. Lu Tzu- Yi et al | 2006 | China | 82 | Sectioning | 54 | 6 | 6 | 10 | 0 | 0 | 0 | 0 | 6† | 18 |
| 9. Awawdeh & Al-Qudah | 2007 | Jordan | 500 | D&C | 58.2 | 4.8 | 1.4 | 14.4 | 16.8 | 0.8 | 1 | 0 | Type 1-3= 1 Type 1-2-3= 0.4 Type 2-1-3= 0.8 Type 2-1-2-1=0.4 | |
| 10. Velmurugan&Sandhya | 2009 | India | 100 | D&C | 72 | 6 | 3 | 10 | 8 | 0 | 0 | 0 | | 1 |
| 11. Jain & Bahuguna | 2011 | India | 138 | D&C | 67.39 | 7.97 | 3.62 | 2.89 | 17.39 | 0.72 | 0 | 0 | | |
| 12. Parekh et al | 2011 | India | 40 | D&C | 50 | 5 | 5 | 25 | 12.5 | 2.5 | 0 | 0 | | |
| 13. Yu et al. | 2012 | China | 174 | CBCT | 86.8 | 0 | 1.7 | 0 | 9.8 | 0 | 0 | 0.6 | | 1.1 |
| 14. Yang et al | 2013 | China | 440 | CBCT | 76.14 | 3.41 | 2.73 | 6.59 | 9.32 | 0 | 0 | 0.68 | | 1.14 |
| 15. Shetty et al | 2014 | India | 118 6 | CBCT | 83.81 | 0.3 | 2.1 | 0.27 | 11.97 | 0.1 | 0 | 0.3 | 0.08‡ | 0.92 |
| 16. Singh and Pawar | 2014 | India | 100 | D&C | 80 | 6 | 0 | 10 | 2 | | | | | |
| 17. Abraham & Gopinath | 2015 | UAE | 100 | Clearing method | 65 | 2 | 3 | 13 | 14 | 0 | 0 | 0 | Type 2-3 = 1% Type 1-4 = 2% | |
| 18. Alfawaz et al | 2019 | Saudi Arabia | 391 | CBCT | 88 | 3.6 | 3.1 | 2 | 1.5 | 0.3 | 0 | 1.5 | | |
| 19. Shrestha et al | 2019 | Nepal | 150 | D&C | 72 | 2.6 | 3.3 | 2.6 | 18.6 | 0 | 0 | 0 | 0.66++ | |
| 20. Present study | 2021 | India | 200 | CBCT | 85.5 | 2.5 | 13 | 0 | 11 | 1 | 3 | 0 | | 1 |

References

- Vertucci F, Seelig A, Gillis R. Root canal morphology of the human maxillary second premolar. Oral Surg Oral Med Oral Pathol 1974;38:456–464.
- Sardar KP, Khokhar NH, Siddiqui I. Frequency of two canals in maxillary second premolar tooth. J Coll Physicians Surg Pakistan 2007;17:12–14.
- **3.** Sert S, Bayirli GS. Evaluation of the root canal configurations of the mandibular and maxillary permanent teeth by gender in the Turkish population. J Endod 2004;30:391–398.
- England MC, Hartwell GR, Lance JR. Detection and treatment of multiple canals in mandibular premolars. J Endod 1991;17:174–178.
- Gulabivala K, Aung TH, Alavi A, Ng YL. Root and canal morphology of Burmese mandibular molars. Int Endod J 2001;34:359–370.
- Martins JNR, Gu Y, Marques D, Francisco H, Caramês J. Differences on the Root and Root Canal Morphologies between Asian and White Ethnic Groups Analyzed by Conebeam Computed Tomography. J Endod 2018;44:1096–1104.
- 7. Slowey RR. Root canal anatomy. Road map to successful endodontics. Dent Clin North Am 1979;23:555–573.
- **8.** Vertucci FJ. Root canal morphology and its relationship to endodontic procedures. Endod Top 2005;10:3–29.
- 9. Yu X, Guo B, Li KZ, et al. Cone-beam computed tomography study of root and canal morphology of mandibular premolars

in a western Chinese population. BMC Med Imaging 2012;12:18.

- **10.** Baisden MK, Kulild JC, Weller RN. Root canal configuration of the mandibular first premolar. J Endod 1992;18:505–508.
- **11.** Cooke HG, Cox FL. C-shaped canal configurations in mandibular molars. J Am Dent Assoc 1979;99:836–839.
- Arayasantiparb R, Banomyong D. Prevalence and morphology of multiple roots, root canals and C-shaped canals in mandibular premolars from cone-beam computed tomography images in a Thai population. J Dent Sci 2021;16:201–207.
- Brea G, Gomez F, Gomez-Sosa JF. Cone-beam computed tomography evaluation of C-shaped root and canal morphology of mandibular premolars. BMC Oral Health 2021;21:1–8.
- **14.** Dou L, Li D, Xu T, Tang Y, Yang D. Root anatomy and canal morphology of mandibular first premolars in a Chinese population. Sci Rep 2017;7:1–7.
- 15. Fan B, Ye W, Xie E, Wu H, Gutmann JL. Three-dimensional morphological analysis of C-shaped canals in mandibular first premolars in a Chinese population. Int Endod J 2012;45:1035–1041.
- Adarsh K, Sharma P, Juneja A. Accuracy and reliability of tooth length measurements on conventional and CBCT images: An in vitro comparative study. J Orthod Sci 2018;7.
- Shetty A, Hegde MN, Tahiliani D, Shetty H, Bhat GT, Shetty S. A Three-Dimensional Study of Variations in Root Canal Morphology Using Cone-Beam Computed Tomography of

Mandibular Premolars in a South Indian Population. J Clin Diagn Res 2014;8:ZC22.

- 18. Jain A, Bahuguna R. Root canal morphology of mandibular first premolar in a gujarati population - an in vitro study. Dent Res J (Isfahan) 2011;8:118–122.
- Velmurugan N, Sandhya R. Root canal morphology of mandibular first premolars in an Indian population: a laboratory study. Int Endod J 2009;42:54–8.
- 20. Abraham SB, Gopinath VK. Root canal anatomy of mandibular first premolars in an Emirati subpopulation: A laboratory study. Eur J Dent 2015;9:476.
- **21.** Vertucci FJ. Root canal anatomy of the human permanent teeth. Oral Surg Oral Med Oral Pathol 1984;58:589–599.
- 22. Zhang W, Tang Y, Liu C, Shen Y, Feng X, Gu Y. Root and root canal variations of the human maxillary and mandibular third molars in a Chinese population: A micro-computed tomographic study. Arch Oral Biol 2018;95:134–140.
- **23.** Trope M, Elfenbein L, Tronstad L. Mandibular premolars with more than one root canal in different race groups. J Endod 1986;12:343–345.
- Cleghorn BM, Christie WH, Dong CCS. The root and root canal morphology of the human mandibular first premolar: a literature review. J Endod 2007;33:509–16.
- Singh S, Pawar M. Root canal morphology of South asian Indian mandibular premolar teeth. J Endod 2014;40:1338– 1341.
- 26. Yang H, Tian C, Li G, Yang L, Han X, Wang Y. A cone-beam computed tomography study of the root canal morphology of mandibular first premolars and the location of root canal orifices and apical foramina in a Chinese subpopulation. J Endod 2013;39:435–438.
- Pineda F, Kuttler Y. Mesiodistal and buccolingual roentgenographic investigation of 7,275 root canals. Oral Surg Oral Med Oral Pathol 1972;33:101–110.

- **28.** Zillich R, Dowson J. Root canal morphology of mandibular first and second premolars. Oral Surg Oral Med Oral Pathol 1973;36:738–744.
- **29.** Çalişkan MK, Pehlivan Y, Sepetçioğlu F, Türkün M, Tuncer SŞ. Root canal morphology of human permanent teeth in a Turkish population. J Endod 1995;21:200–204.
- **30.** Lu TY, Yang SF, Pai SF. Complicated Root Canal Morphology of Mandibular First Premolar in a Chinese Population Using the Cross Section Method. J Endod 2006;32:932–6.
- **31.** Awawdeh LA, Al-Qudah AA. Root form and canal morphology of mandibular premolars in a Jordanian population. Int Endod J 2008;41:240–248.
- **32.** Parekh V, Shah N, Joshi H. Root canal morphology and variations of mandibular premolars by clearing technique: an in vitro study. J Contemp Dent Pract 2011;12:318–321.
- 33. Alfawaz H, Alqedairi A, Al-Dahman YH, et al. Evaluation of root canal morphology of mandibular premolars in a Saudi population using cone beam computed tomography: A retrospective study. Saudi Dent J 2019;31:137–142.
- **34.** Shrestha R, Srii R, Shrestha D. Diversity of root canal morphology in mandibular first premolar. Kathmandu Univ Med J 2019;17:223–228.
- 35. Habib AA, Kalaji MN, Al Saysd TJ, Al Jawfi KA. Root canal configurations of the first and second mandibular premolars in the population of north Syria. J Taibah Univ Med Sci 2015;10:391–395.
- 36. Xu J, Shao MY, Pan HY, et al. A proposal for using contralateral teeth to provide well-balanced experimental groups for endodontic studies. Int Endod J 2016;49:1001–8.
- Johnsen GF, Dara S, Asjad S, Sunde PT, Haugen HJ. Anatomic Comparison of Contralateral Premolars. J Endod 2017;43:956–963.