P-ISSN: 2204-1990; E-ISSN: 1323-6903 DOI: 10.47750/cibg.2021.27.02.309

Knowledge, Attitude And Practice Regarding Nickel Hypersensitivity In Patients Undergoing Orthodontic Therapy Among Interns. - A Questionnaire Survey.

SANDHYA.A¹, REMMIYA MARY VARGHESE^{2*}, SENTHIL MURUGAN.P³

¹Saveetha Dental College and Hospitals, Saveetha Institute of Medical and Technical Sciences (SIMATS) Saveetha University, Chennai, India

Email: 151501040.sdc@saveetha.com¹, remmiyav.sdc@saveetha.com², Senthilmuruganp.sdc@saveetha.com³

Abstract: Nickel is one of the most common causes of allergic contact dermatitis and produces more allergic reactions than all other metals combined. Currently, several brands of orthodontic wires are made of nickel titanium alloy and potentially have a high enough nickel content to provoke manifestations of allergic reactions in the oral cavity. The main objective of this study is to evaluate the knowledge and awareness on nickel hypersensitivity in patients among interns in a private dental college. The study was conducted in a private dental college, Chennai. The study group consisted of the interns and postgraduate students. Total number of studies included in the study was 200. The statistical analysis was done using SPSS software (SPSS version 21.0, SPSS, Chicago II, USA). The data was analysed using a chi-square test. The p value of less than 0.05 was considered to be statistically significant. About 73% of them are aware about Nickel Hypersensitivity and 74.5% are aware that Nickel hypersensitivity is one of the most common causes for allergic contact dermatitis. Higher number of females reported to know about Nickel Hypersensitivity when compared to males. There was no significant difference between the gender and awareness about Nickel Hypersensitivity. Chi- square value: 0.059, p value: 0.807. Dental students in this study have a good level of knowledge and positive attitude towards nickel hypersensitivity in patients. However, the knowledge acquired must be implemented in their daily practice and provide the better treatment required for the patients. A better knowledge of nickel hypersensitivity is essential for safe practice in dentistry. This will ensure the provision of better and safer dental health care services for the population.

Keywords: Interns; Nickel dermatitis; Nickel dermatitis; Orthodontic appliances; Skin lesion.

INTRODUCTION

An allergic response is one in which certain components of the immune system react excessively to a foreign substance. Two key allergic reactions have been described in the literature. Type I hypersensitivity reactions are an immediate antibody mediated allergic response, occurring within minutes or hours after direct skin or mucosal contact with the allergen. (Poley and Slater, 2000) Nickel alloys are widely used in the orthodontic in brackets, wires, bands and other orthodontic accessories. Nickel allergy occurs more frequently than allergy to all other metals combined.(Lowey, 1993) It is estimated that 11% of all women and 20% of women between the ages of 16 and 35 years have a sensitivity to nickel.6-8 Nickel-induced contact dermatitis is a Type IV delayed hypersensitivity immune response occurring at least 24 hours after exposure.(Al-Tawil, Marcusson and Moller, 1985)(Loon et al., 1988) It has been shown that the level of nickel in saliva and serum increases significantly after the insertion of fixed orthodontic appliances. (Souza, de Souza and de Menezes, 2008) Nickel leaching from orthodontic bands, brackets, stainless steel or Ni-Ti archwires has been shown in vitro to occur within the first week and then decline thereafter.(Barrett, Bishara and Quinn, 1993) It is suggested that a threshold concentration of approximately 30 ppm of nickel may be sufficient to elicit a cytotoxic response. (Bour et al., 1994) Scientific evidence suggests that orthodontic treatment is not associated with increase of Ni hypersensitivity, unless patients have a history of previous exposure to Ni. People with cutaneous piercing are considered a significant risk factor for Ni allergy. (Thyssen et al., 2007) however, oral exposure to nickel through dental braces prior to ear piercing reduces the risk of developing nickel allergy. (Mortz et al., 2002)(Hoogstraten et al., 2008)

²Senior lecturer, Department of Orthodontic dentistry, Saveetha Dental college and Hospitals, Saveetha institute of medical and technical sciences (SIMATS), Saveetha University, Chennai, India.

³Associate Professor, Department of Oral and Maxillofacial Surgery, Saveetha Dental college and Hospitals Saveetha institute of medical and technical sciences (SIMATS), Saveetha University, Chennai, India *Corresponding Author:

Signs and symptoms of nickel allergy includes gingivitis, gingival hyperplasia, lip desquamation, burning sensation in the mouth, metallic taste, angular cheilitis, and periodontitis. (Staerkjaer and Menne, 1990) (Bishara, Barrett and Selim, 1993) In chronic cases, the affected mucosa is typically in contact with the causal agent and appears erythematous or hyperkeratotic to ulcerated. (Genelhu et al., 2005) Extraoral manifestations of nickel allergy may have an intra oral origin. (Schultz et al., 2004) If a nickel allergy is still in question, a diagnosis can be confirmed by a dermatologist by conducting a cutaneous sensitivity test called a patch test (Table 1) using 5% nickel sulphate in petroleum jelly. (Menné et al., 1987)

If intra - oral signs and symptoms are present and a diagnosis of nickel hypersensitivity is established, the fixed or removable prosthesis should be replaced with another nickel free alloy. The nickel titanium archwires should be removed and replaced with a stainless steel archwire which is low in nickel content or preferably a titanium molybdenum alloy (TMA) arch wire, known as "TMA", which does not contain nickel. Most patients who develop a reaction to Ni-Ti archwires subsequently tolerate stainless steel without a reaction. (Toms, 1988) Other options include fibre reinforced composite wires, Gold plated wires, Ion-implanted nickel-titanium arch wires or Plastic/Resin-coated nickel-titanium arch wires. (Kucukyildirim and Eker, 2012)

Stainless steel brackets are generally considered safe. However, nickel free alternative brackets to stainless steel include Ceramic brackets produced using polycrystalline alumina, single crystal sapphire, and zirconia, Polycarbonate brackets, Titanium brackets and Gold plated brackets and plastic brackets in selected cases Fixed appliances may be substituted with plastic aligners. Extra-oral metal components, including metal studs in headgear, are of greatest concern due to greater sensitivity of skin. Plastic coated headgear studs may be a better alternative for such patients. Previously our team has conducted numerous clinical trials, few review papers and surveys (Kumar et al., 2011; Felicita, Shanthasundari and Chandrasekar, 2012; Dinesh and Saravana Dinesh, 2013; Jain, 2014; Kamisetty, 2015; Krishnan, 2015; Rubika, Sumathi Felicita and Sivambiga, 2015; Viswanath et al., 2015; Sivamurthy and Sundari, 2016; Felicita and Sumathi Felicita, 2017a, 2017b, 2018; Samantha, 2017; Vikram and Raj Vikram, 2017; Krishnan, Pandian and Kumar, 2018). Our team has rich experience in research and we have collaborated with numerous authors over various topics in the past decade (Deogade, Gupta and Ariga, 2018; Ezhilarasan, 2018; Ezhilarasan, Sokal and Najimi, 2018; Jeevanandan and Govindaraju, 2018; J et al., 2018; Menon et al., 2018; Prabakar et al., 2018; Rajeshkumar et al., 2018, 2019; Vishnu Prasad et al., 2018; Wahab et al., 2018; Dua et al., 2019; Duraisamy et al., 2019; Ezhilarasan, Apoorva and Ashok Vardhan, 2019; Gheena and Ezhilarasan, 2019; Malli Sureshbabu et al., 2019; Mehta et al., 2019; Panchal, Jeevanandan and Subramanian, 2019; Rajendran et al., 2019; Ramakrishnan, Dhanalakshmi and Subramanian, 2019; Sharma et al., 2019; Varghese, Ramesh and Veeraiyan, 2019; Gomathi et al., 2020; Samuel, Acharya and Rao, 2020)

The present study aims to evaluate interns' awareness on nickel hypersensitivity in patients reporting to the clinic.

MATERIALS AND METHOD

Study design:

A cross sectional questionnaire based study was carried out among dental students of a Private Dental College in chennai who are practising in clinics.

Sample:

This study was conducted in an online setting. The sample comprised 200 participants. Simple random sampling methodology was employed.

Approval:

Informed consent and ethical approval was obtained from the Institutional Ethical Committee and Scientific Review Board of the University. (SDC/SIHEC/2020/DIASDATA/0619-0320).

Questionnaire:

The questionnaire was framed with the help of experts in the field. The questionnaire kept the study group in mind and questions were linked to curriculum content of nickel hypersensitivity. A self – administrated questionnaire consisting of 10 close ended questions. The dental students answered the questionnaire through an online setting survey planet. There were 10 questions to assess their knowledge, attitude and to judge their practices of the respondents on nickel hypersensitivity patients reporting to the clinic.

Statistical analysis:

The data from their response were entered in the excel sheets. The data was later exported to SPSS Software (SPSS version 21.0, SPSS, Chicago II, USA) for statistical analysis. The data was analysed using a chi-square test. The p value of less than 0.05 was considered to be statistically significant.

RESULTS AND DISCUSSION

Out of 200 participants, about 73% of them are aware about nickel hypersensitivity [GRAPH 1] and 74.5% are aware that nickel hypersensitivity is one of the most common causes for allergic contact dermatitis [GRAPH 2]. About 55% of the students gave positive responses about nickel being one of the components used in making

orthodontic wires [GRAPH 3]. From the chart it is observed that 38% of dental students have said that the daily dietary requirements for humans is 25 - 35 and 30 - 40. There was an equal distribution among these two [GRAPH 4]. Only about 12% were aware that 300 - 600 being the average diet supply of nickel to the human body per day[GRAPH 5]. About 94% of them were aware about the threshold concentration requirement of nickel to cause allergic reactions[GRAPH 6]. It is observed that 75.50% of them were aware about the intraoral signs and symptoms of nickel hypersensitivity[GRAPH 7]. Only about 19% were about the longevity of symptoms of nickel hypersensitivity[GRAPH 8]. About 48.5% of students were aware that patch test is the test used to diagnose nickel hypersensitivity in patients[GRAPH 9]. 70.5% of students are aware that both fibre reinforced composite wire and TMA are the alternative used in dentistry with various other alternatives[GRAPH 10]. Higher number of females reported to know about Nickel Hypersensitivity when compared to males. There was no significant difference between the gender and awareness about Nickel Hypersensitivity.(p value: 0.807 (p>0.05 statistically non significant))[GRAPH 11] About 41.50% females reported to know about Nickel Hypersensitivity when compared to males. There was no significant difference between the gender and awareness about Nickel Hypersensitivity being one of the common causes for allergic contact dermatitis.(p value: 0.886(p>0.05 statistically non significant))[GRAPH 12] About 29.50% of females reported to know about Nickel being one of the components used in making orthodontic appliances when compared to males. There was no significant difference between the gender and awareness about Nickel being one of the components used in making orthodontic appliances. (p value: 0.595(p>0.05 statistically non significant))[GRAPH 13] About 22.50% females reported to know about the daily dietary requirement of Nickel for humans when compared to males. There was no significant difference between the gender and awareness about the daily dietary requirement of Nickel for humans. (p value: 0.729(p>0.05 statistically non significant))[GRAPH 14] About 20% females reported to know about the average diet supply of Nickel for humans per day when compared to males. There was no significant difference between the gender and awareness about the average diet supply of Nickel for humans per day. (p value: 0.882(p>0.05 statistically non significant))[GRAPH 15] About 26% females know about the threshold concentration requirement of Nickel when compared to males. There was no significant difference between the gender and awareness about the threshold concentration requirement of Nickel. (p value: 0.811(p>0.05 statistically non significant))[GRAPH 16] About 42% females know about the intraoral signs and symptoms of Nickel Hypersensitivity when compared to males. There was no significant difference between the gender and awareness about the intraoral signs and symptoms of Nickel Hypersensitivity. (p value: 0.859(p>0.05 statistically non significant))[GRAPH 17] About 22% females know about the longevity of the symptoms of Nickel Hypersensitivity when compared to males. There was no significant difference between the gender and awareness about the longevity of the symptoms of Nickel Hypersensitivity.(p value: 0.902 (p>0.05 statistically non significant))[GRAPH 18] About 24.50% females know about the test used to diagnose Nickel Hypersensitivity when compared to males. There was significant difference between the gender and awareness about the test used to diagnose Nickel Hypersensitivity. (p value: 0.034 (p<0.05 statistically significant))[GRAPH 19] About 41% females know about the alternative used for Nickel containing stainless steel wire when compared to males. There was no significant difference between the gender and about the alternative used for Nickel containing stainless steel wire (p value: 0.607(p>0.05 statistically non significant))[GRAPH 20]

From this study, 73% of them are aware about nickel hypersensitivity and it is one of the most common causes for allergic contact dermatitis. However, a Scandinavian authors noted that a considerable number of nickelsensitive patients developed dermatitis flare-up at sites different from those that have been in direct contact with nickel-plated items, so they speculated an endogenous exposure to Ni in patients with hip prosthesis or dental alloys.(Christensen and Möoller, 1975; Frigerio et al., 2011; Yilmaz, Ozdemir and Yilmaz, 2012)

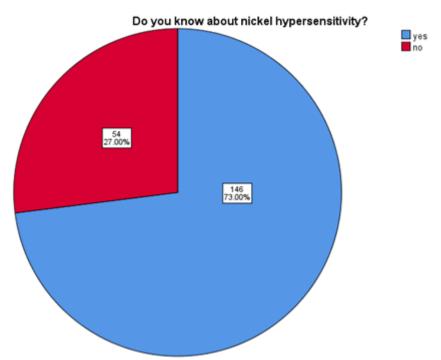
From this study, 55% of the students gave positive responses about nickel being one of the components used in making orthodontic wires. Orthodontic arch wires that contained up to 54% nickel caused no cytotoxic effect. Similarly, it has been determined that the maximum amount of nickel released from orthodontic arch wires was 700 times lower than the amount necessary to elicit cytotoxic reaction in a human peripheral blood mononuclear cell culture. These findings are important as the use of high level nickel containing 'shape-memory' wires is increasing in Orthodontics.(Grimsdottir, Hensten-Pettersen and Kullmann, 1992; Jia et al., 1999)

There is rising concern about the biocompatibility of dental materials; this might be due to a real increase in the occurrence of allergic reactions to the materials or to an increase in awareness of adverse effects from these materials.3 Allergy in patients undergoing orthodontic treatment can be seen due to several reasons and these include nickel allergy, allergy to the acrylic resins that are used during treatment. (Kucukyildirim and Eker, 2012)

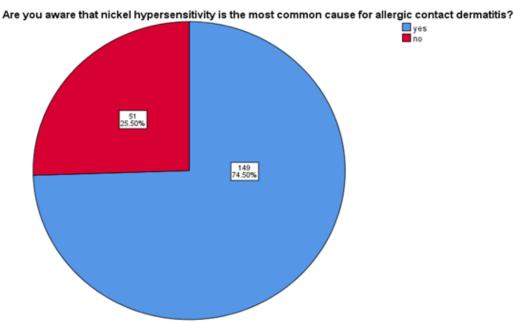
Safe and effective practice depends on identifying patients with allergy along with knowledge of materials that can potentially cause them. Orthodontists should have basic understanding of allergic reactions and should be efficient enough to manage them.

It is important for a clinician to not only know the physical and mechanical properties of the materials being used, but also of the biologic compatibility of the material. Knowledge of alternatives to allergy causing materials is also of prime importance in efficient management of patients in routine clinical practice.

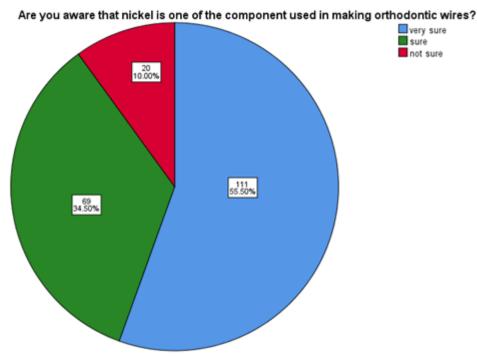
A detailed history, with special attention to previous allergic reactions, is the main prognostic factor to avoid allergic reactions during orthodontic therapy. The clinician should be mindful of these reactions during the course of orthodontic treatment, and should know to diagnose and subsequent action to be taken in the treatment plan.



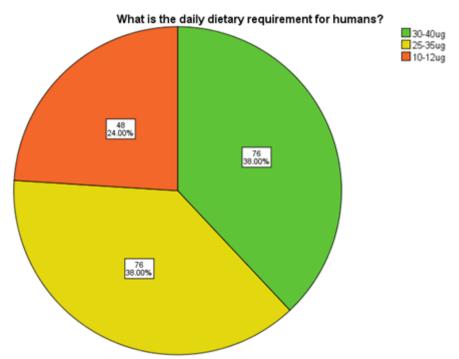
Graph 1: The pie chart showing distribution of study population who were aware of Nickel hypersensitivity. About 73% know about Nickel hypersensitivity and 54% gave negative responses



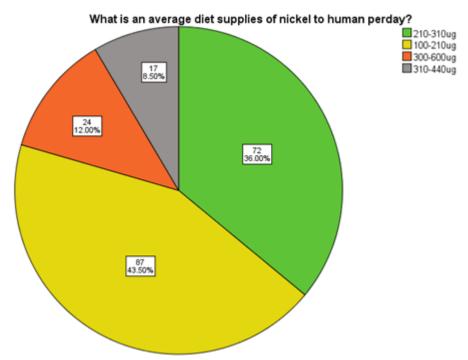
Graph 2: The pie chart showing distribution of study population who were aware that Nickel hypersensitivity is one of the causes for allergic contact dermatitis. About 74.5% of students gave positive responses.



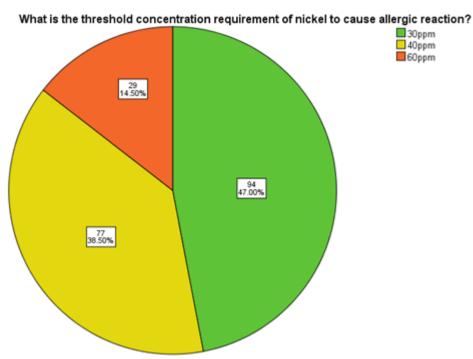
Graph 3: The pie chart showing distribution of study population who were aware of Nickel being one of the components used to make orthodontic wires. About 55.50% of them gave positive responses.



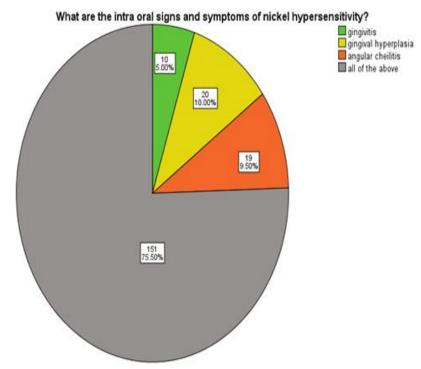
Graph 4: The pie chart showing distribution of study population who were asked about the daily dietary requirements of nickel for humans. About 38% of them answered both 30-40 ug and 25-35 ug being the daily dietary requirements for humans.



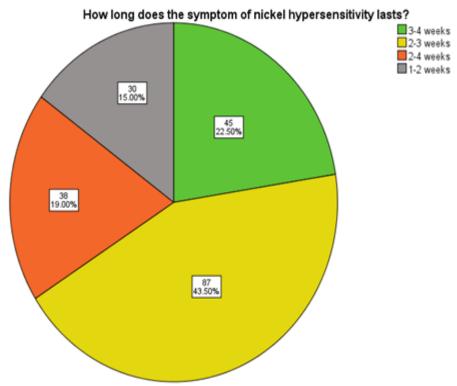
Graph 5: The pie chart showing distribution of study population who were asked about the average diet supply of nickel to humans per day. Only about 12% of students gave positive responses



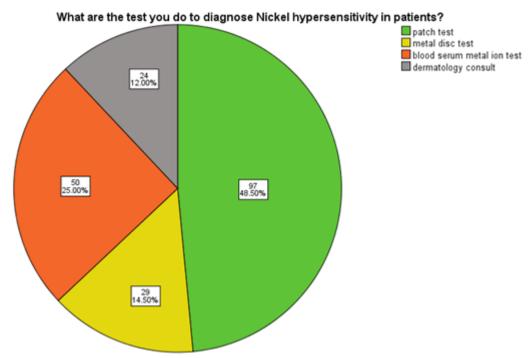
Graph 6: The pie chart showing distribution of study population who were asked about the threshold concentration of Nickel to cause allergic reactions. About 94% of students gave positive responses.



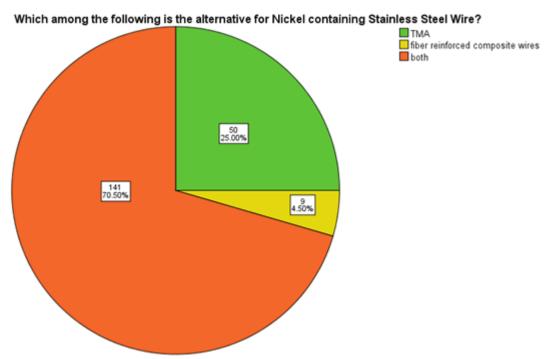
Graph 7: The pie chart showing distribution of study population who were aware of the intra oral symptoms of Nickel hypersensitivity. About 75.50% of students know about the intra oral symptoms of Nickel hypersensitivity.



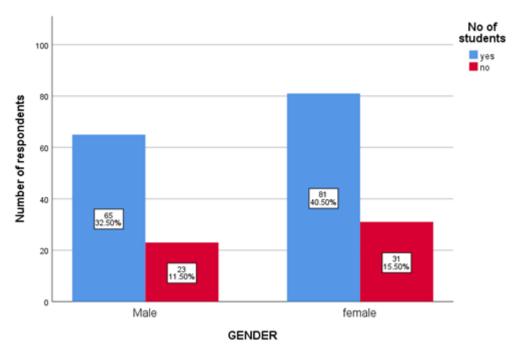
Graph 8: The pie chart showing distribution of study population who were asked about the duration of Nickel hypersensitivity symptoms to last. Only about 19% of the students gave positive responses.



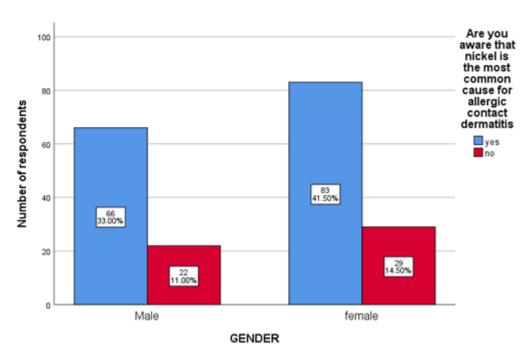
Graph 9: The pie chart showing distribution of study population who were aware about the test used to diagnose Nickel hypersensitivity. About 48.50% know about the diagnostic test for Nickel hypersensitivity.



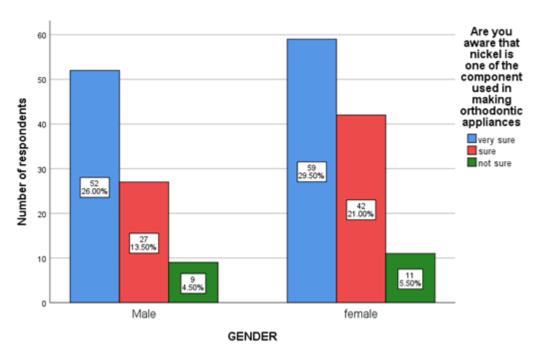
Graph 10: The pie chart showing distribution of study population who were aware about the alternative for Nickel containing stainless steel wire. About 70.50% of the students gave positive responses.



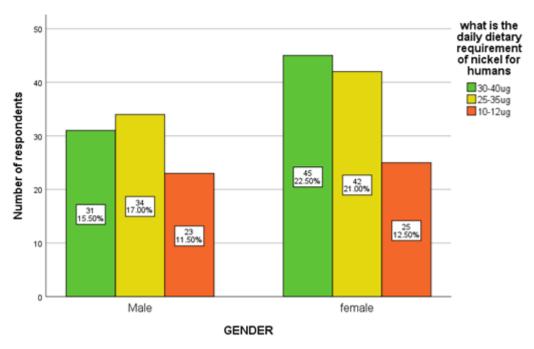
Graph 11: The bar graph showing the association between gender and awareness about Nickel Hypersensitivity. Higher number of females reported to know about Nickel Hypersensitivity when compared to males. There was no significant difference between the gender and awareness about Nickel Hypersensitivity. (Chi – Square, p value: 0.807 (p>0.05 statistically non significant))



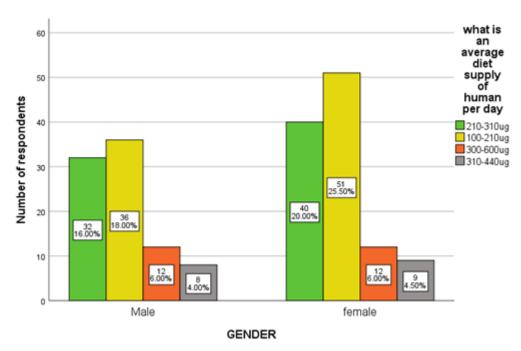
Graph 12: The bar graph showing the association between gender and awareness about Nickel Hypersensitivity being one of the common causes for allergic contact dermatitis. It is observed that a higher number of females reported to know about Nickel Hypersensitivity when compared to males. There was no significant difference between the gender and awareness about Nickel Hypersensitivity being one of the common causes for allergic contact dermatitis. (Chi – Square, p value: 0.886(p>0.05 statistically non significant))



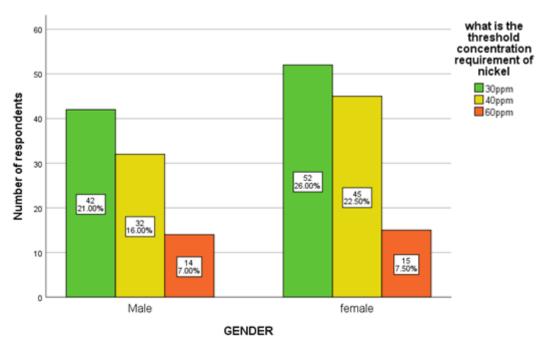
Graph 13: The bar graph showing the association of gender and awareness about Nickel being one of the components used in making orthodontic appliances. It is observed that a higher number of females reported to know about Nickel being one of the components used in making orthodontic appliances when compared to males. There was no significant difference between the gender and awareness about Nickel being one of the components used in making orthodontic appliances. (Chi – Square, p value: 0.595(p>0.05 statistically non significant))



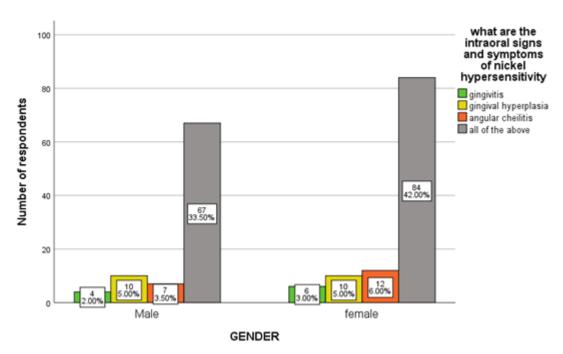
Graph 14: The bar graph showing the association of gender and awareness about the daily dietary requirement of Nickel for humans. It is observed that a higher number of females reported to know about the daily dietary requirement of Nickel for humans when compared to males. There was no significant difference between the gender and awareness about the daily dietary requirement of Nickel for humans. (Chi – Square, p value: 0.729(p>0.05 statistically non significant))



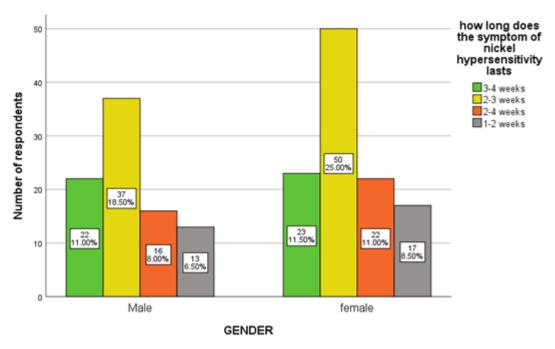
Graph 15: The bar graph showing the association of gender and awareness about the average diet supply of Nickel for humans per day. It is observed that a higher number of females reported to know about the average diet supply of Nickel for humans per day when compared to males. There was no significant difference between the gender and awareness about the average diet supply of Nickel for humans per day. (Chi – Square, p value: 0.882(p>0.05 statistically non significant))



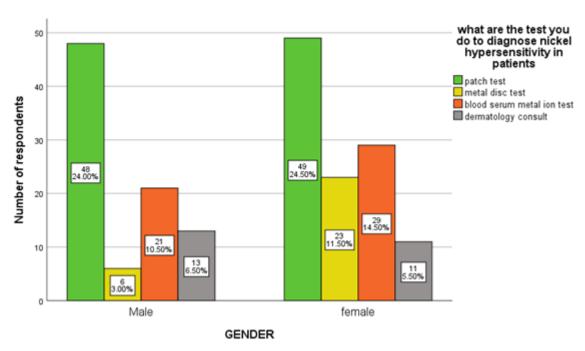
Graph 16: The bar graph showing the association of gender and awareness about the threshold concentration requirement of Nickel. It is observed that a higher number of females know about the threshold concentration requirement of Nickel when compared to males. There was no significant difference between the gender and awareness about the threshold concentration requirement of Nickel. (Chi – Square, p value: 0.811(p>0.05 statistically non significant))



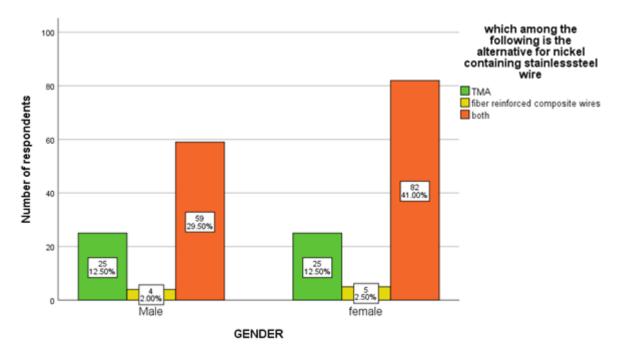
Graph 17: The bar graph showing the association of gender and awareness about the intraoral signs and symptoms of Nickel Hypersensitivity. It is observed that a higher number of females know about the intraoral signs and symptoms of Nickel Hypersensitivity when compared to males. There was no significant difference between the gender and awareness about the intraoral signs and symptoms of Nickel Hypersensitivity. (Chi – Square, p value: 0.859(p>0.05 statistically non significant))



Graph 18: The bar graph showing the association of gender and awareness about the longevity of the symptoms of Nickel Hypersensitivity. It is observed that a higher number of females know about the longevity of the symptoms of Nickel Hypersensitivity when compared to males. There was no significant difference between the gender and awareness about the longevity of the symptoms of Nickel Hypersensitivity. (Chi – Square, p value: 0.902 (p>0.05 statistically non significant))



Graph 19: The bar graph showing the association of gender and awareness about the test used to diagnose Nickel Hypersensitivity. It is observed that a higher number of females know about the test used to diagnose Nickel Hypersensitivity when compared to males. There was significant difference between the gender and awareness about the test used to diagnose Nickel Hypersensitivity. (Chi – Square, p value: 0.034 (p<0.05 statistically significant))



Graph 20: The bar graph showing the association of gender and awareness about the alternative used for Nickel containing stainless steel wire. It is observed that Higher number of females know about the alternative used for Nickel containing stainless steel wire when compared to males. There was no significant difference between the gender and about the alternative used for Nickel containing stainless steel wire (Chi – Square, p value: 0.607(p>0.05 statistically non significant))

Sandhya.A et al/ Knowledge, Attitude And Practice Regarding Nickel Hypersensitivity In Patients Undergoing Orthodontic Therapy Among Interns. - A Questionnaire Survey.

Our institution is passionate about high quality evidence based research and has excelled in various fields ((Pc, Marimuthu and Devadoss, 2018; Ramesh et al., 2018; Ezhilarasan, Apoorva and Ashok Vardhan, 2019; Ramadurai et al., 2019; Sridharan et al., 2019; Vijayashree Priyadharsini, 2019; Mathew et al., 2020)

LIMITATIONS

The main limitations of this study was limited sample size and it was confined to limited geographical population. For further scope of the study increased sample size with inclusion of varied population ethnicity would give better results.

CONCLUSION

Dental students in this study have a good level of knowledge and positive attitude towards nickel hypersensitivity in patients. However, the knowledge acquired must be implemented in their daily practice and provide the better treatment required for the patients. Continuing education programs and refreshing courses regarding nickel hypersensitivity in patients are necessary to update the knowledge of dental practitioners.

AUTHORS CONTRIBUTION

First author, sandhya performed the data collection by reviewing patient details, filtering required data, analysing and interpreting statistics and contributed to manuscript writing.

Second author, Dr. Remmiya Mary Varghese contributed to conception of study title, study design, analysed the collected data, statistics and interpretation and also critically revised the manuscript.

Third author, Dr. Senthil Murugan P participated in the study and revised the manuscript. All the three authors have discussed the results and contributed to the final manuscript.

ACKNOWLEDGEMENT

This research was supported by saveetha dental college and hospital. We thank the department of Orthodontics, Saveetha Dental College for providing insight and expertise that greatly assisted this research.

CONFLICT OF INTEREST

The authors declare that there is no conflict of interests.

REFERENCE

- 1. Al-Tawil, N. G., Marcusson, J. A. and Moller, E. (1985) 'T and B Lymphocytes in Patients with Nickel Sensitivity', Scandinavian Journal of Immunology, pp. 495–502. doi: 10.1111/j.1365-3083.1985.tb01908.x.
- 2. Barrett, R. D., Bishara, S. E. and Quinn, J. K. (1993) 'Biodegradation of orthodontic appliances. Part I. Biodegradation of nickel and chromium in vitro', American Journal of Orthodontics and Dentofacial Orthopedics, pp. 8–14. doi: 10.1016/0889-5406(93)70098-9.
- 3. Bishara, S. E., Barrett, R. D. and Selim, M. I. (1993) 'Biodegradation of orthodontic appliances. Part II. Changes in the blood level of nickel', American Journal of Orthodontics and Dentofacial Orthopedics, pp. 115–119. doi: 10.1016/s0889-5406(05)81760-3.
- 4. Bour, H. et al. (1994) 'Establishment of Nickel-Specific T Cell Lines from Patients with Allergic Contact Dermatitis: Comparison of Different Protocols', Clinical Immunology and Immunopathology, pp. 142–145. doi: 10.1006/clin.1994.1180.
- 5. Christensen, O. B. and Möoller, H. (1975) 'External and internal exposure to the antigen in the hand eczema of nickel allergy', Contact Dermatitis, pp. 136–141. doi: 10.1111/j.1600-0536.1975.tb05354.x.
- 6. Deogade, S., Gupta, P. and Ariga, P. (2018) 'Effect of monopoly-coating agent on the surface roughness of a tissue conditioner subjected to cleansing and disinfection: A Contact Profilometric In vitro study', Contemporary Clinical Dentistry, p. 122. doi: 10.4103/ccd.ccd_112_18.
- 7. Dinesh, S. P. S. and Saravana Dinesh, S. P. (2013) 'An Indigenously Designed Apparatus for Measuring Orthodontic Force', JOURNAL OF CLINICAL AND DIAGNOSTIC RESEARCH. doi: 10.7860/jcdr/2013/7143.3631.
- 8. Dua, K. et al. (2019) 'The potential of siRNA based drug delivery in respiratory disorders: Recent advances and progress', Drug development research, 80(6), pp. 714–730.
- 9. Duraisamy, R. et al. (2019) 'Compatibility of Nonoriginal Abutments With Implants: Evaluation of Microgap at the Implant-Abutment Interface, With Original and Nonoriginal Abutments', Implant dentistry, 28(3), pp. 289–295.
- 10. Ezhilarasan, D. (2018) 'Oxidative stress is bane in chronic liver diseases: Clinical and experimental perspective', Arab journal of gastroenterology: the official publication of the Pan-Arab Association of Gastroenterology, 19(2), pp. 56–64.
- 11. Ezhilarasan, D., Apoorva, V. S. and Ashok Vardhan, N. (2019) 'Syzygium cumini extract induced reactive oxygen species-mediated apoptosis in human oral squamous carcinoma cells', Journal of oral pathology &

- medicine: official publication of the International Association of Oral Pathologists and the American Academy of Oral Pathology, 48(2), pp. 115–121.
- 12. Ezhilarasan, D., Sokal, E. and Najimi, M. (2018) 'Hepatic fibrosis: It is time to go with hepatic stellate cell-specific therapeutic targets', Hepatobiliary & pancreatic diseases international: HBPD INT, 17(3), pp. 192–197.
- 13. Felicita, A., Shanthasundari, K. K. and Chandrasekar, S. (2012) 'Determination of craniofacial relation among the subethnic Indian population: A modified approach (Sagittal relation)', Indian Journal of Dental Research, p. 305. doi: 10.4103/0970-9290.102210.
- 14. Felicita, A. S. and Sumathi Felicita, A. (2017a) 'Orthodontic management of a dilacerated central incisor and partially impacted canine with unilateral extraction A case report', The Saudi Dental Journal, pp. 185–193. doi: 10.1016/j.sdentj.2017.04.001.
- 15. Felicita, A. S. and Sumathi Felicita, A. (2017b) 'Quantification of intrusive/retraction force and moment generated during en-masse retraction of maxillary anterior teeth using mini-implants: A conceptual approach', Dental Press Journal of Orthodontics, pp. 47–55. doi: 10.1590/2177-6709.22.5.047-055.oar.
- 16. Felicita, A. S. and Sumathi Felicita, A. (2018) 'Orthodontic extrusion of Ellis Class VIII fracture of maxillary lateral incisor The sling shot method', The Saudi Dental Journal, pp. 265–269. doi: 10.1016/j.sdentj.2018.05.001.
- 17. Frigerio, E. et al. (2011) 'Metal sensitivity in patients with orthopaedic implants: a prospective study', Contact Dermatitis, pp. 273–279. doi: 10.1111/j.1600-0536.2011.01886.x.
- 18. Genelhu, M. C. L. S. et al. (2005) 'Characterization of nickel-induced allergic contact stomatitis associated with fixed orthodontic appliances', American Journal of Orthodontics and Dentofacial Orthopedics, pp. 378–381. doi: 10.1016/j.ajodo.2005.03.002.
- 19. Gheena, S. and Ezhilarasan, D. (2019) 'Syringic acid triggers reactive oxygen species-mediated cytotoxicity in HepG2 cells', Human & experimental toxicology, 38(6), pp. 694–702.
- 20. Gomathi, A. C. et al. (2020) 'Anticancer activity of silver nanoparticles synthesized using aqueous fruit shell extract of Tamarindus indica on MCF-7 human breast cancer cell line', Journal of Drug Delivery Science and Technology, p. 101376. doi: 10.1016/j.jddst.2019.101376.
- 21. Grimsdottir, M. R., Hensten-Pettersen, A. and Kullmann, A. (1992) 'Cytotoxic effect of orthodontic appliances', The European Journal of Orthodontics, pp. 47–53. doi: 10.1093/ejo/14.1.47.
- 22. Hoogstraten, I. M. W. et al. (2008) 'Reduced frequency of nickel allergy upon oral nickel contact at an early age', Clinical & Experimental Immunology, pp. 441–445. doi: 10.1111/j.1365-2249.1991.tb05746.x.
- 23. Jain, R. K. (2014) 'Comparison of Intrusion Effects on Maxillary Incisors Among Mini Implant Anchorage, J-Hook Headgear and Utility Arch', JOURNAL OF CLINICAL AND DIAGNOSTIC RESEARCH. doi: 10.7860/jcdr/2014/8339.4554.
- 24. Jeevanandan, G. and Govindaraju, L. (2018) 'Clinical comparison of Kedo-S paediatric rotary files vs manual instrumentation for root canal preparation in primary molars: a double blinded randomised clinical trial', European Archives of Paediatric Dentistry, pp. 273–278. doi: 10.1007/s40368-018-0356-6.
- 25. Jia, W. et al. (1999) 'Nickel release from orthodontic arch wires and cellular immune response to various nickel concentrations', Journal of Biomedical Materials Research, pp. 488–495. doi: 3.0.co;2-d">10.1002/(sici)1097-4636(1999)48:4<488::aid-jbm14>3.0.co;2-d.
- 26. J, P. C. et al. (2018) 'Prevalence and measurement of anterior loop of the mandibular canal using CBCT: A cross sectional study', Clinical implant dentistry and related research, 20(4), pp. 531–534.
- 27. Kamisetty, S. K. (2015) 'SBS vs Inhouse Recycling Methods-An Invitro Evaluation', JOURNAL OF CLINICAL AND DIAGNOSTIC RESEARCH. doi: 10.7860/jcdr/2015/13865.6432.
- 28. Krishnan, S. (2015) 'Effect of Bisphosphonates on Orthodontic Tooth Movement—An Update', JOURNAL OF CLINICAL AND DIAGNOSTIC RESEARCH. doi: 10.7860/jcdr/2015/11162.5769.
- 29. Krishnan, S., Pandian, K. and Kumar, S. (2018) 'Angular photogrammetric analysis of the soft-tissue facial profile of Indian adults', Indian Journal of Dental Research, p. 137. doi: 10.4103/ijdr.ijdr_496_16.
- 30. Kucukyildirim, B. O. and Eker, A. A. (2012) 'Surface Roughness Changes and Corrosion on Nickel Titanium Orthodontic Wires Compared to Stainless Steel Wires in Various Artificial Salivas', Materials Testing, pp. 261–265. doi: 10.3139/120.110327.
- 31. Kumar, K. R. R. et al. (2011) 'Depth of resin penetration into enamel with 3 types of enamel conditioning methods: A confocal microscopic study', American Journal of Orthodontics and Dentofacial Orthopedics, pp. 479–485. doi: 10.1016/j.ajodo.2010.10.022.
- 32. Loon, L. A. J. et al. (1988) 'T-lymphocyte and Langerhans cell distribution in normal and allergically induced oral mucosa in contact with nickel-containing dental alloys', Journal of Oral Pathology and Medicine, pp. 129–137. doi: 10.1111/j.1600-0714.1988.tb01899.x.
- 33. Lowey, M. N. (1993) 'Allergic contact dermatitis associated with the use of an Interlandi headgear in a patient with a history of atopy', British Dental Journal, pp. 67–72. doi: 10.1038/sj.bdj.4808226.
- 34. Malli Sureshbabu, N. et al. (2019) 'Concentrated Growth Factors as an Ingenious Biomaterial in

- Regeneration of Bony Defects after Periapical Surgery: A Report of Two Cases', Case reports in dentistry, 2019, p. 7046203.
- 35. Mathew, M. G. et al. (2020) 'Evaluation of adhesion of Streptococcus mutans, plaque accumulation on zirconia and stainless steel crowns, and surrounding gingival inflammation in primary ...', Clinical oral investigations. Available at: https://link.springer.com/article/10.1007/s00784-020-03204-9.
- 36. Mehta, M. et al. (2019) 'Oligonucleotide therapy: An emerging focus area for drug delivery in chronic inflammatory respiratory diseases', Chemico-biological interactions, 308, pp. 206–215.
- 37. Menné, T. et al. (1987) 'Patch test reactivity to nickel alloys', Contact Dermatitis, pp. 255–259. doi: 10.1111/j.1600-0536.1987.tb01448.x.
- 38. Menon, S. et al. (2018) 'Selenium nanoparticles: A potent chemotherapeutic agent and an elucidation of its mechanism', Colloids and Surfaces B: Biointerfaces, pp. 280–292. doi: 10.1016/j.colsurfb.2018.06.006.
- 39. Mortz, C. G. et al. (2002) 'Nickel Sensitization in Adolescents and Association with Ear Piercing, Use of Dental Braces and Hand Eczema', Acta Dermato-Venereologica, pp. 359–364. doi: 10.1080/000155502320624096.
- 40. Panchal, V., Jeevanandan, G. and Subramanian, E. M. G. (2019) 'Comparison of post-operative pain after root canal instrumentation with hand K-files, H-files and rotary Kedo-S files in primary teeth: a randomised clinical trial', European archives of paediatric dentistry: official journal of the European Academy of Paediatric Dentistry, 20(5), pp. 467–472.
- 41. Pc, J., Marimuthu, T. and Devadoss, P. (2018) 'Prevalence and measurement of anterior loop of the mandibular canal using CBCT: A cross sectional study', Clinical implant dentistry and related research. Available at: https://europepmc.org/article/med/29624863.
- 42. Poley, G. E. and Slater, J. E. (2000) 'Latex allergy', Journal of Allergy and Clinical Immunology, pp. 1054–1062. doi: 10.1067/mai.2000.106925.
- 43. Prabakar, J. et al. (2018) 'Comparative Evaluation of Retention, Cariostatic Effect and Discoloration of Conventional and Hydrophilic Sealants A Single Blinded Randomized Split Mouth Clinical Trial', Contemporary clinical dentistry, 9(Suppl 2), pp. S233–S239.
- 44. Rajendran, R. et al. (2019) 'Comparative Evaluation of Remineralizing Potential of a Paste Containing Bioactive Glass and a Topical Cream Containing Casein Phosphopeptide-Amorphous Calcium Phosphate: An in Vitro Study', Pesquisa Brasileira em Odontopediatria e Clínica Integrada, pp. 1–10. doi: 10.4034/pboci.2019.191.61.
- 45. Rajeshkumar, S. et al. (2018) 'Biosynthesis of zinc oxide nanoparticles using Mangifera indica leaves and evaluation of their antioxidant and cytotoxic properties in lung cancer (A549) cells', Enzyme and microbial technology, 117, pp. 91–95.
- 46. Rajeshkumar, S. et al. (2019) 'Antibacterial and antioxidant potential of biosynthesized copper nanoparticles mediated through Cissus arnotiana plant extract', Journal of photochemistry and photobiology. B, Biology, 197, p. 111531.
- 47. Ramadurai, N. et al. (2019) 'Effectiveness of 2% Articaine as an anesthetic agent in children: randomized controlled trial', Clinical oral investigations, 23(9), pp. 3543–3550.
- 48. Ramakrishnan, M., Dhanalakshmi, R. and Subramanian, E. M. G. (2019) 'Survival rate of different fixed posterior space maintainers used in Paediatric Dentistry A systematic review', The Saudi dental journal, 31(2), pp. 165–172.
- 49. Ramesh, A. et al. (2018) 'Comparative estimation of sulfiredoxin levels between chronic periodontitis and healthy patients A case-control study', Journal of periodontology, 89(10), pp. 1241–1248.
- 50. Rubika, J., Sumathi Felicita, A. and Sivambiga, V. (2015) 'Gonial Angle as an Indicator for the Prediction of Growth Pattern', World Journal of Dentistry, pp. 161–163. doi: 10.5005/jp-journals-10015-1334.
- 51. Samantha, C. (2017) 'Comparative Evaluation of Two Bis-GMA Based Orthodontic Bonding Adhesives A Randomized Clinical Trial', JOURNAL OF CLINICAL AND DIAGNOSTIC RESEARCH. doi: 10.7860/jcdr/2017/16716.9665.
- 52. Samuel, S. R., Acharya, S. and Rao, J. C. (2020) 'School Interventions-based Prevention of Early-Childhood Caries among 3-5-year-old children from very low socioeconomic status: Two-year randomized trial', Journal of public health dentistry, 80(1), pp. 51–60.
- 53. Schultz, J. C. et al. (2004) 'Cutaneous and Oral Eruption from Oral Exposure to Nickel in Dental Braces', Dermatitis (formerly American Journal of Contact Dermatitis), p. 154. doi: 10.2310/6620.2004.04022.
- 54. Sharma, P. et al. (2019) 'Emerging trends in the novel drug delivery approaches for the treatment of lung cancer', Chemico-biological interactions, 309, p. 108720.
- 55. Sivamurthy, G. and Sundari, S. (2016) 'Stress distribution patterns at mini-implant site during retraction and intrusion—a three-dimensional finite element study', Progress in Orthodontics. doi: 10.1186/s40510-016-0117-1.
- 56. Souza, R. M. de, de Souza, R. M. and de Menezes, L. M. (2008) 'Nickel, Chromium and Iron Levels in the Saliva of Patients with Simulated Fixed Orthodontic Appliances', The Angle Orthodontist, pp. 345–350.

- doi: 10.2319/111806-466.1.
- 57. Sridharan, G. et al. (2019) 'Evaluation of salivary metabolomics in oral leukoplakia and oral squamous cell carcinoma', Journal of oral pathology & medicine: official publication of the International Association of Oral Pathologists and the American Academy of Oral Pathology, 48(4), pp. 299–306.
- 58. Staerkjaer, L. and Menne, T. (1990) 'Nickel allergy and orthodontic treatment', The European Journal of Orthodontics, pp. 284–289. doi: 10.1093/ejo/12.3.284.
- 59. Thyssen, J. P. et al. (2007) 'The epidemiology of contact allergy in the general population prevalence and main findings', Contact Dermatitis, pp. 287–299. doi: 10.1111/j.1600-0536.2007.01220.x.
- 60. Toms, A. P. (1988) 'The corrosion of orthodontic wire', The European Journal of Orthodontics, pp. 87–97. doi: 10.1093/ejo/10.2.87.
- 61. Varghese, S. S., Ramesh, A. and Veeraiyan, D. N. (2019) 'Blended Module-Based Teaching in Biostatistics and Research Methodology: A Retrospective Study with Postgraduate Dental Students', Journal of dental education, 83(4), pp. 445–450.
- 62. Vijayashree Priyadharsini, J. (2019) 'In silico validation of the non-antibiotic drugs acetaminophen and ibuprofen as antibacterial agents against red complex pathogens', Journal of periodontology, 90(12), pp. 1441–1448.
- 63. Vikram, N. R. and Raj Vikram, N. (2017) 'Ball Headed Mini Implant', JOURNAL OF CLINICAL AND DIAGNOSTIC RESEARCH. doi: 10.7860/jcdr/2017/24358.9240.
- 64. Vishnu Prasad, S. et al. (2018) 'Report on oral health status and treatment needs of 5-15 years old children with sensory deficits in Chennai, India', Special care in dentistry: official publication of the American Association of Hospital Dentists, the Academy of Dentistry for the Handicapped, and the American Society for Geriatric Dentistry, 38(1), pp. 58–59.
- 65. Viswanath, A. et al. (2015) 'Obstructive sleep apnea: awakening the hidden truth', Nigerian journal of clinical practice, 18(1), pp. 1–7.
- 66. Wahab, P. U. A. et al. (2018) 'Scalpel Versus Diathermy in Wound Healing After Mucosal Incisions: A Split-Mouth Study', Journal of oral and maxillofacial surgery: official journal of the American Association of Oral and Maxillofacial Surgeons, 76(6), pp. 1160–1164.
- 67. Yilmaz, A., Ozdemir, C. E. and Yilmaz, Y. (2012) 'A Delayed Hypersensitivity Reaction to a Stainless Steel Crown', Journal of Clinical Pediatric Dentistry, pp. 235–238. doi: 10.17796/jcpd.36.3.d1327wn32361u04n.