MEDICAL SCIENCE

To Cite:

Lath T, Rathi N, Pawar M, Bendgude V, Sinnarkar S, Mehta V, Noor T. Knowledge, attitude, and practices amongst pediatric dentists in Pune towards Local Anaesthetic allergy and test dose in dental operatory: A questionnaire study. *Medical Science* 2023; 27: e294ms3060. doi: https://doi.org/10.54905/disssi/v27i137/e294ms3060

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Peer-Review History

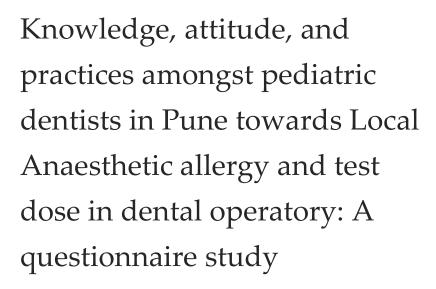
Received: 11 May 2023 Reviewed & Revised: 15/May/2023 to 07/July/2023 Accepted: 11 July 2023 Published: 18 July 2023

Peer-review Method

External peer-review was done through double-blind method.

Medical Science pISSN 2321–7359; eISSN 2321–7367

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ABSTRACT

Aim: The aim of the survey is evaluate the current knowledge, attitude and practices of paediatric dentist towards local anaesthesia allergy and test dose administration in the dental operatory. Material and Methods: A 25-question questionnaire was framed in accordance with local anesthesia guidelines amongst 103 pediatric dentists residing in Pune. Data collection was done by sending the questionnaire to the participants through google forms via emails/WhatsApp. The responses to questionnaires were summarized and this was accomplished by converting the online recorded information into representative numbers (codes). The data was entered into the Statistical Package for Social Sciences (SPSS) and analyzed. Results: Most of them, about 62.83% were not well aware of protocols for LA allergy testing. Maximum of participants, 30% did not know if the dose of epinephrine (1:1000) can be given in a child less than 30 kgs. Epinephrine was the drug chosen by 88% to be their first choice in management of anaphylaxis. 70% preferred Intramuscular as initial route for epinephrine injection. 60% said they have never seen a patient with local adverse reaction after use of local anesthesia. Conclusion: The lack of information among dental practitioners about maximum doses and dose estimates for local anesthetics for children was alarming; mainly as LA systemic toxicity was dose dependent. Our study suggested conducting additional educational courses to keep both general dental practitioners and specialists up-to-date on proper application of these vital components of dentistry.

Keywords: Pediatric dentists, knowledge, practice, local anesthetic, allergy, test dose



1. INTRODUCTION

The child patient's most prevalent fear in the dental office is the fear of pain, one which nature has endowed the child as a means of self-protection. Unfortunately, most children are conditioned to expect pain as a certain part of dentistry's routine – even before they experience any dental service themselves (Litt, 1996). Local anesthetics (LA) are a reliable, effective, low-cost, and widely available anesthetic. These medications are grouped as esters or amides, and their effect might be short, intermediate, or long. Other constituents, like vasoconstrictors, are equally important in LA delivery and the local anesthetic agent in the cartridge (Wahl and Brown, 2010). Many factors influence the required dose of LA, which involved age and weight, and medically impaired children must be considered when estimating the maximum dose of LA or a vasoconstrictor (VC).

Patients who experience systemic reactions after receiving LA are frequently referred to as 'allergic' to the local anesthetic. Drug overdose, vasovagal reactions, fast absorption, cardiovascular, intravascular injection, psychogenic or idiosyncratic reactions and central nervous system, were the most prominent causes for systemic sequelae (Sambrook et al., 2011). Although it is frequently claimed by patients, a true allergic reaction to dental LA is uncommon, including publications claiming allergic reactions to LA accounted for 1% of all adverse systemic reactions only (Sambrook et al., 2011; Haas, 2002).

A detailed understanding of these reactions, on the other hand, is beneficial because it gives the patient the highest chance of recovery and future usage of LA for pain management. Metabisulphite, the preservative added to several formulations, is the most likely allergen among the components included in LA cartridge (Sambrook et al., 2011). It is added as an antioxidant in products containing adrenaline, a VC (Haas, 2002). A case report of a verified allergy to ethylenediaminetetraacetic acid (EDTA) from LA was recently published in the literature (Russo et al., 2014).

Typically, skin prick and intradermal tests of suspected allergen triggers are used to identify allergies. On the basis of such skin tests, LA-induced allergic responses have been evaluated earlier (Seymour et al., 1999; Bhole et al., 2012). LA are regarded as relatively safe, but the high frequency of injections provided makes unpleasant reactions unavoidable (Haas, 2002). There has been a notable increase in the availability of local anesthesia (LA) products, including over-the-counter medications, in recent times. Consequently, it is possible that there has been a corresponding rise in the number of individuals who have developed sensitivity or an allergic response to LA.

In other countries, evaluations of dentists' knowledge and attitudes on LA-induced anaphylaxis have been conducted (Thyssen et al., 2008; Hepner and Castells, 2003; Lieberman et al., 2006). These investigations showed that dentists lacked sufficient skill in the identification and treatment of such responses. In India, the Pune region has not yet been the subject of such research. Thus, the present study aimed to assess the level of knowledge, attitude, and practice among pediatric dentists in Pune towards local anesthesia allergy and test dose in the dental operatory.

2. MATERIALS AND METHODS

A cross-sectional closed survey was performed amongst 103 Pediatric dentists who were practicing in the city of Pune, Maharashtra, India over a period of 6 months, from August 2022 to January 2023 which was conducted after acquiring the scientific and ethical clearance from the institutional review, scientific and ethical committee from Dr DY Patil Dental College & Hospital, Pune (DYPDCH/IEC/164/151/20). Since this was an e-survey, it was not possible to collect separate informed consents. Therefore, a brief outline of the purpose of study along with the inclusion criteria was explained and that their participation was voluntary keeping their details confidential. Hence, the response received from the Pediatric dentists was considered as an implied consent. The details of their demographics, knowledge, attitude and practice were collected through a pre-designed questionnaire by the principal investigator through an e-survey using Google forms (Table 1).

A 25-question questionnaire was framed in accordance with local anesthesia guidelines. Questions were later cross categorized based on knowledge, attitude and practice. Relatively easy questions were placed at the beginning of the questionnaire with difficult questions placed at the end. Data collection was done by sending the questionnaire to the participants through google forms via e-mails/WhatsApp. Before starting the study, the examiner pilot tested the online format of the questionnaire amongst 10 participants.

We wanted to be able to have confidence in the findings of the study, so we made sure that the questionnaire, when it was correctly filled out and administered, measured what it claimed it would measure consistently. In a nutshell, the questionnaire met both the criteria for validity and reliability. For face validity, the questionnaire had a professional appearance so that there was better chance of eliciting serious responses. In order to ensure that the questionnaire contained content validity, it was given to 10

Subject Matter Experts. They marked the questions as essential or non-essential based on the relevance, clarity, brevity of the questions. The data was collected and analyzed for Content validity ratio (CVR).

The reliability of the questionnaire, also known as its reproducibility, determines whether or not it operates consistently. The "test-retest method of reliability" was utilized, which refers to the ability of the questionnaire to produce comparable findings when it is given to the same individual on two distinct times. The correlation between the results is positively correlated with the reliability of the questionnaire. Ten participants took part in the test - retest reliability. Reliability of the questionnaire was checked using SPSS software. Kappa value was found to be 0.9 which indicated the questionnaire to be highly reliable. The online recorded data was transformed into representative numbers (codes), which allowed for the summary of the questionnaire replies.

The coding system was considered when the questionnaire was being developed. SPSS Version 26 software package (SPSS inc., IBM, and Chicago, IL, USA) was used to analyze the data. Numbers and Percentages were calculated for each item. Chi-squared test was applied to compare knowledge, attitude and practice based on the demographic variables (Number of years of experience in Pediatric dentists).

3. RESULTS

The present study was undertaken to assess the knowledge, attitude and practice of 103 Pediatric dentists residing in Pune city, with respect to local anesthesia hypersensitivity and test dose in dental operatory.

Table 1 Participants' demographic information

Duration of practice		
Less than 5 years	19	18.4%
5-10 years	75	72.9%
More than 10 years	9	8.7%

72.9% of the participants had practiced for 5 to 10 years, about 18.4% participants had practiced for less than 5 years and only 8.7% had experience of more than 10 years.

Table 2 Knowledge regarding Local Anaesthetic (LA) allergy and test dose in dental operatory

Questions	Frequency N=103	Percent (%)	
1. What types of adverse reactions are commonly associated with local			
anaesthesia administration?			
Systemic toxicity	12	11.6%	
Local toxicity	12	11.6%	
Psychogenic reactions	12	11.6%	
Paraesthesia	12	11.6%	
Soft tissue injury	13	12.9%	
Allergic reactions	13	12.9%	
All of the above	17	16.2%	
Don't Know	12	11.6%	
2. What components of local anaesthesia solution may cause			
hypersensitivity?			
Local anaesthesia	21	21.3%	
Preservatives	60	58.2%	
Anti-oxidants	3	2.2%	
Adrenaline	4	3.8%	
Don't know	15	14.5%	
3. Hypersensitivity reactions are more common in			
Ester group LA	60	58.2%	
Amide group LA	30	29.1%	
None of the above	0	0	

Don't know	13	12.7%	
4. Are you well aware of protocols for LA allergy testing?			
Yes	60	58.2%	
No	43	41.8%	
5. Types of hypersensitivity reactions that may occur with LA			
Type 1 or Anaphylactic reactions	70	67.9%	
Type 2 or Cytotoxic reactions	4	4%	
Type 3 or Immune complex reactions	4	4%	
Type 4 or T cell-mediated reactions	12	12%	
All of the above	11	11%	
Don't know	2	1.1%	
6. Allergic reaction to LA can be seen w	rithin		
<2 hours	61	59%	
<6 hours	2	2 %	
6-24 hours	17	16%	
Over 24 hours	3	3%	
Any of the above	20	20%	
7. What dose of epinephrine (1:1000) ca	n be given in a child l	less than 30	
kgs?			
0.10 mg/day	21	20 %	
0.15 mg /day	23	22%	
0.20 mg/day	20	20%	
0.30 mg/day	8	8%	
Don't know	31	30%	

Most of the participants, 66.95% selected soft tissue injury as adverse reactions are commonly associated with local anesthesia administration. About 58.2% of them were unaware of the procedures for LA allergy testing. 67.9% described Type 1 or Anaphylactic reactions to be the hypersensitivity reactions that may occur with LA and 59% said allergic reaction to LA can be seen within <2 hours. Maximum of participants, 30% did not know if the dose of epinephrine (1:1000) can be given in a child less than 30 kgs.

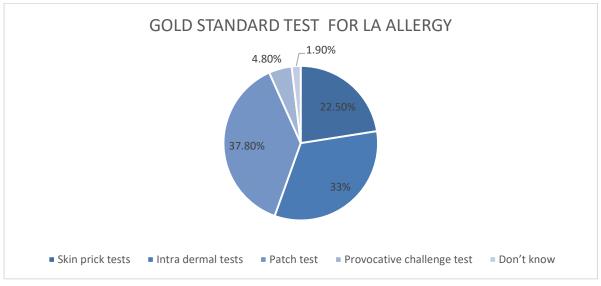


Figure 1 Knowledge regarding Gold standard for testing LA allergy

Figure 1 show only 4.8% of participants consider Provocative Challenge Test (PCTs) as the gold standard for testing LA allergy while 37.80%, 33% and 22.50% think patch test, intradermal tests and skin prick test as the standard test respectively.

Table 3 Attitude regarding Local Anaesthetic (LA) allergy and test dose in dental operatory

Questions	Frequency N=103	Percent (%)
When a patient reports a suspected allergy to local anesth		. ,
I do not begin the treatment	7	7%
I make a skin prick test with the suspected drug by myself	20	19%
I refer the patient to an allergy specialist with the		
suspected drug for testing	71	69%
I treat the patient without local anaesthesia	5	5%
2. Which route, do you prefer giving the test dose in?		
Intra oral	7	7%
Intra dermal	96	93%
Others	0	0%
3. Would you do an allergy test for patients without a prior	suggestive history?	
Yes	26	25%
Not at all	34	33%
Sometimes if necessary	43	42%
4. What response, following the test dosage, do you think w	vould be indicative of	anaphylaxis?
Skin rashes	31	30%
Itching	3	3%
Dyspnoea	4	3.5%
Sudden fainting	4	3.5%
All of the above	61	60%
Not sure	0	0%
5. What medication would you use first to treat anaphylaxis	s?	
Epinephrine	91	88%
Antihistamine	7	7%
Corticosteroids	1	0.9%
Glucagon	0	0%
Salbutamol	1	1.1%
Oxygen	3	3%
Don't know	0	0%
6. Which approach do you favour for epinephrine injection	as a first course of act	tion?
Intramuscular	72	70%
Subcutaneous	21	20%
Intravenous	4	4.5%
I'm not sure	6	5.5%
7. Do you think it is important to keep oxygen and respirate	ory masks in the oper	atory for
emergency management?		
Yes	103	100%
No	0	0%

The attitude of operators about LA allergy and test dose is in (Table 3). 93% of participants, or almost all, preferred administering the test dosage intradermally. 88% of people selected epinephrine as their preferred medication option for treating anaphylaxis. 70% prefer intramuscular injections of epinephrine as the first approach. Every participant, that is 100%, though it is important to keep oxygen and respiratory masks in the operatory for emergency management.

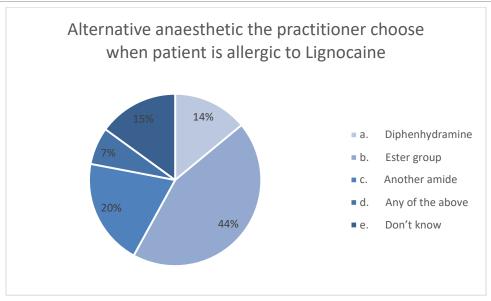


Figure 2 Attitude of operators about choice of an alternative for lignocaine

Figure 2 shows the knowledge of operators about an alternative for lignocaine. 44% participants would opt for an ester group LA if patient is allergic to Lignocaine.

Table 4 Practice regarding Local Anaesthetic (LA) allergy and test dose in dental operatory

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Questions	Frequency N=103	Percent (%)	
1. Have you encountered a patient who experienced a			
local adverse response following local anaesthesia?			
Yes	41	40%	
No	62	60%	
2. Have you encountered a patient who experienced a			
systemic adverse response following local anaesthesia?			
Yes	11	10.5%	
No	92	89.5%	
3. Do you maintain the following drugs at your office?			
Epinephrine	67	65%	
Antihistamine	23	22%	
Corticosteroids	6	6%	
Glucagon	2	2%	
Salbutamol	4	4%	
None of the above	1	1%	

Table 4 shows the response from dental operatory regarding LA allergy and test dose. In the survey, 60% of respondents said they had never seen a patient have a local adverse reaction and 89.5% of respondents said they had never seen a patient have any systemic response after receiving local anesthesia.

Figure 3 shows the responses from operators about seeking history of LA administration. 83% do not give a test dose with each appointment while sometimes they administer a test dose. However, most of them take previous history of LA administration and any associated food allergy.

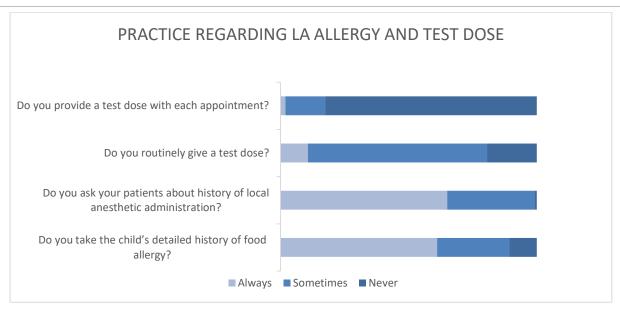


Figure 3 Practice regarding LA allergy and test dose in the operatory

4. DISCUSSION

True allergy to LA, particularly amide local anesthetics, is extremely rare (Sambrook et al., 2011). Rood, (2000) performed cutaneous and intraoral challenges on 44 adult patients who were thought to have a LA allergy and came to the conclusion that most of the symptoms were psychogenic in nature and that no allergy was ever observed to be present. Only one of these patients over this time had a documented allergy to mepivacaine, demonstrating a rather infrequent reaction to dental local anesthetics (Gall et al., 1996).

In developed nations, anaphylactic responses to LA following dental procedures have been observed, with a frequency of 1 in 3,500 to 1 in 13,000 instances (Vervloet et al., 1999; Laxenaire, 1999). Australia and Norway have recorded incidences of 1 in 10,000 to 1 in 20,000 and 1 in 6,000, respectively, in recent studies (Hepner and Castells, 2003; Lieberman et al., 2006; Webb and Lieberman, 2006). Therefore, the purpose of the current study was to evaluate the degree of knowledge, attitude, and practice among pediatric dentists in Pune about local anesthesia allergy and test dosage in dental operatory. A total of 103 members participated in the present study. Pediatric dentist residing in Pune city were approached for the study.

According to studies, the majority of practicing dentists utilize low dosages of local anesthetic and vasoconstrictor (Carolina et al., 2015). Anaphylactic drug allergy is one of the most unpredictable ADRs because it is unrelated to dosage and can be lethal. Hypersensitivity to local anesthetic solution can sometimes be misinterpreted by practitioners as toxicity to LA agent and/or the VC, as well as anxiety symptoms (Canfield and Gage, 1987). The British Society for Allergy and Clinical Immunology's Standards of Care Committee explained how to investigate suspected anaphylaxis during general anesthesia. It is noted that skin prick tests and intradermal testing have not been validated due to the rarity of a true LA allergy. As a result, it is recommended to proceed to progressive subcutaneous challenge as needed/appropriate only after such testing (Ewan et al., 2010).

Approximately 5-10% of all adverse drug reactions (ADRs) are classified as drug allergies, which fall under the category of unpredictable ADRs. Drug allergies comprise a wide range of immunologically induced hypersensitivity reactions with diverse mechanisms and clinical manifestations (Dewachter et al., 2009; Sicherer and Leung, 2013). Allergies to preservatives in local anesthetic solutions are substantially more common (though still uncommon). Preservatives, such as bisulfite, which is included in multidose vials, are another unusual source of an immediate-type reaction to LA (Pavlidakey et al., 2009).

Atopic illnesses, like hay fever, allergic rhinitis, and asthma, were not risk factors for LA sensitivity, but dentists frequently direct patients with atopic diseases to allergy clinics for drug allergy testing. Our study reported that 58.2% practitioners believed preservatives to be the cause (Table 2 Q2). Anaphylaxis is one of the most critical clinical settings seen in everyday medicine. In over 70% of patients, symptoms ranging from the skin to the CVS and respiratory systems are present at the same time. Anaphylaxis must be identified and treated as soon as possible (Lieberman, 2006) because anaphylaxis may be encountered in their ordinary practice, albeit it is not common (Sampson et al., 2006). Our study reported 67.9% practitioners were aware about this type I allergy (Table 2 Q5).

The most successful method of managing drug allergies is the avoidance of the offending substance or discontinuing its use. Whenever feasible, alternative pharmaceuticals with entirely different chemical structures should be chosen as substitutes for the original medication. When selecting alternative agents, it is crucial to consider the potential for cross-reactivity between different medications. Similar to another study, our findings showed that most of the dentists are aware of this fact and 58.2% believed that Ester group was most commonly associated with Hypersensitivity reactions (Cetinkaya et al., 2011) (Table 2 Q3).

The ester group of anesthetics carries a higher risk of unfavorable allergic responses than the amide group does. The preservatives methyl paraben and metabisulfite were most likely the cause of some of the adverse responses that persons described. Caine, the anesthetic agent, did not cause allergic reactions in all of the people who reported allergic reactions. Epinephrine, antihistamines, and corticosteroids are all relatively inexpensive medications, and as a result, they should be stocked at every medical facility (Anchor and Settipane, 2004).

Our study findings were in agreement with other research showing about 65% of the dentists kept epinephrine ready in their offices and 88% administered it in ADRs (Table 4 Q3). When faced with an anaphylactic episode, the recommended course of treatment involves administering an intramuscular injection of epinephrine into the lateral thigh (Simons and Simons, 2010). The indicated approach of administering epinephrine via the IM method was similar to that of the previous studies, with around 70% of dentists preferring the IM route for epinephrine administration (Table 3 Q6).

The dose of epinephrine prescribed is determined by the patient's weight. For children and adults weighing 30 kg or more, the average dose is 0.3 mg (Lee and Vadas, 2011). A maximum of practitioners of our study, 30% were not aware of the dosage while many were divided on their opinion. The other options like antihistamines and corticosteroids are also reported but in minute numbers by the practitioners in our study. Corticosteroids and antihistamines given systemically are other therapy alternatives for severe systemic reactions; however, they should never be given before or in place of epinephrine when dealing with anaphylactic patients (Sheikh, 2013; Sheikh et al., 2007; Kaira and Dabral, 2014).

Many complications can arise from the incorrect administration of LA; thus, various protocols need to be followed when employing LA of which only 43.26% of the general dental practitioners were aware of while the remaining 62.83% of the respondents were unaware (Liau et al., 2008; Conrado et al., 2007; Rishiraj et al., 2005). It is particularly critical when anesthetics are delivered to medically impaired patients or children by an unaware practitioner, because the doses of local anesthetics and vasoconstrictors administered to these patients and children differ from those administered to healthy individuals (Conrado et al., 2007; American Academy on Pediatric Dentistry Council on Clinical Affairs, 2008; Lipp et al., 1993; Mochizuki et al., 1989).

Lastly, it is exceedingly rare to experience allergic reactions to local anesthesia; however, in the event they do occur, they can pose life-threatening risks. The lifetime prevalence of anaphylaxis is increasing, and it is noteworthy that the incidence of anaphylaxis in children is higher than in adults (Lee et al., 2020). Patients with anaphylaxis often lack awareness of the seriousness of their condition when visiting their dentists. Therefore, it is essential for dental professionals to be well-versed in the causes and management of anaphylaxis. Furthermore, nationwide continuing dental education (CDE) programs focused on anaphylaxis, such as conferences, workshops, journal clubs, and seminars, could prove beneficial in extensively educating dentists on this matter.

5. CONCLUSION

Anaphylaxis occurring during dental procedures is infrequent; however, when it does happen, the consequences can be severe. The concerning lack of understanding among dental professionals regarding optimal dosage limits and estimations for local anesthetics in children, as observed in our study, should be corroborated to the fact that systemic toxicity from local anesthetics is dependent on the dosage administered. This suggests the need for additional educational programs aimed at keeping dentists, both general practitioners and specialists, updated on the correct utilization of these crucial elements in dentistry.

The study's findings highlight the dentists' limited experience in managing such reactions. To tackle this issue, it is crucial to enhance dentists' proficiency in basic life support. Attending continuing dental education courses in this field, encompassing workshops and practical training, should be deemed necessary.

Acknowledgement

We thank the participants who were all contributed samples to the study.

Author Contributions

Details of contribution of each authors regards manuscript work & production.

Ethical approval

The study was approved by the Medical Ethics Committee of Dr DY Patil Dental College & Hospital, Pune (Ethical approval code: DYPDCH/IEC/164/151/20).

Informed consent

Informed consent was obtained from all individual participants as implied consents who chose to participate in the study.

Funding

This study has not received any external funding.

Conflict of interest

The authors declare that there is no conflict of interests.

Data and materials availability

All data sets collected during this study are available upon reasonable request from the corresponding author.

REFERENCES AND NOTES

- American Academy on Pediatric Dentistry Council on Clinical Affairs. Guideline on appropriate use of local anesthesia for pediatric dental patients. Pediatr Dent 2008; 3 0(7 Suppl):134-9.
- Anchor J, Settipane RA. Appropriate use of epinephrine in anaphylaxis. Am J Emerg Med 2004; 22(6):488-90. doi: 10.10 16/j.ajem.2004.07.016
- Bhole MV, Manson AL, Seneviratne SL, Misbah SA. IgE-mediated allergy to local anaesthetics: Separating fact from perception: A UK perspective. Br J Anaesth 2012; 108(6):903 e11. doi: 10.1093/bja/aes162
- Canfield DW, Gage TW. A guideline to local anaesthetic allergy testing. Anaesth Prog 1987; 34(5):157–163.
- Carolina A, Franca J, Soares M, Salles L. Dentist's knowledge regarding signs and symptoms of the systemic toxicity of local anesthetic solutions. Rev Gauch Odontol 2015; 63(1):41–46. doi: 10.1590/1981-863720150001000062962
- Cetinkaya F, Sezgin G, Aslan OM. Dentists' knowledge about anaphylaxis caused by local anaesthetics. Allergol Immunopathol (Madr) 2011; 39(4):228-31. doi: 10.1016/j.aller .2 010.07.009
- Conrado VC, De-Andrade J, De-Angelis GA, De-Andrade AC, Timerman L, Andrade MM, Moreira DR, Sousa AG, Sousa JE, Piegas LS. Cardiovascular effects of local anesthesia with vasoconstrictor during dental extraction in coronary patients. Arq Bras Cardiol 2007; 88(5):507-13. doi: 10.1590/s0066-782x2007000500002
- 8. Dewachter P, Mouton-Faivre C, Emala CW. Anaphylaxis and anesthesia: Controversies and new insights. Anesthesio logy 2009; 111:1141-50. doi: 10.1097/ALN.0b013e3181bbd443
- Ewan PW, Dugué P, Mirakian R, Dixon TA, Harper JN, Nasser SM; BSACI. BSACI guidelines for the investigation of suspected anaphylaxis during general anaesthesia. Clin

- Exp Allergy 2010; 40(1):15-31. doi: 10.1111/j.1365-2222.2009. 03404.x
- Gall H, Kaufmann R, Kalveram CM. Adverse reactions to local anesthetics: Analysis of 197 cases. J Allergy Clin Immunol 1996; 97(4):933-7. doi: 10.1016/s0091-6749(96)80067
- 11. Haas DA. An update on local anaesthetics in dentistry. J Can Dent Assoc 2002; 68(9):546–551.
- 12. Hepner DL, Castells MC. Anaphylaxis during the perioperative period. Anesth Analg 2003; 97(5):1381–95. doi: 10.1213/01.ANE.0000082993.84883.7D
- 13. Kaira LS, Dabral E. A survey to access knowledge and practice among dentists regarding local anesthetic dosage in three cities of Uttarakhand. Eur J Gen Dent 2014; 3:105-8.
- 14. Laxenaire MC. Again, and always allergy. Ann Fr Anesth Reanim 1999; 18(8):831–833. doi: 10.1016/s0750-7658(00)8818 9-7
- Lee JK, Vadas P. Anaphylaxis: Mechanisms and management. Clin Exp Allergy 2011; 41(7):923-38. doi: 10.11 11/j.1365-2222.2011.03779.x
- Lee WS, An J, Jung YH, Jee HM, Chae KY, Park YA, Han MY, Lee KS. Characteristics and Treatment of Anaphylaxis in Children Visiting a Pediatric Emergency Department in Korea. Biomed Res Int 2020; 2020:2014104. doi: 10.1155/2020 /2014104
- 17. Liau FL, Kok SH, Lee JJ, Kuo RC, Hwang CR, Yang PJ, Lin CP, Kuo YS, Chang HH. Cardiovascular influence of dental anxiety during local anesthesia for tooth extraction. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2008; 105(1): 16-26. doi: 10.1016/j.tripleo.2007.03.015
- 18. Lieberman P, Camargo CA, Bohlke K, Jick H, Miller RL, Sheikh A, Simons FE. Epidemiology of anaphylaxis findings of the American College of Allergy, Asthma and

- Immunology, Epidemiology of Anaphylaxis Working Group. Ann Allergy Asthma Immunol 2006; 97:596–602. doi: 10.1016/S1081-1206(10)61086-1
- Lieberman P. Anaphylaxis. Med Clin North Am 2006; 90(1): 77-95. doi: 10.1016/j.mcna.2005.08.007
- Lipp M, Dick W, Daubländer M, Fuder H, Stanton-Hicks M. Exogenous and endogenous plasma levels of epinephrine during dental treatment under local anesthesia. Reg Anesth 1993; 18(1):6-12.
- 21. Litt MD. A model of pain and anxiety associated with acute stressors: Distress in dental procedures. Behav Res Ther 1996; 34(5-6):459-76. doi: 10.1016/0005-7967(96)00015-0
- 22. Mochizuki M, Yokota S, Murata Y, Watanabe H, Nishibori M, Suzuki N, Nishibori M, Kubota Y. Changes in heart rate and blood pressure during dental procedures with local anesthesia. Anesth Prog 1989; 36(4-5):234-5.
- 23. Pavlidakey PG, Brodell EE, Helms SE. Diphenhydramine as an alternative local anesthetic agent. J Clin Aesthet Dermatol 2009; 2(10):37-40.
- 24. Rishiraj B, Epstein JB, Fine D, Nabi S, Wade NK. Permanent vision loss in one eye following administration of local anesthesia for a dental extraction. Int J Oral Maxillofac Surg 2005; 34(2):220-3. doi: 10.1016/j.ijom.2004.04.010
- 25. Rood JP. Adverse reaction to dental local anaesthetic injection 'allergy' is not the cause. Br Dent J 2000; 189(7):38 0–84. doi: 10.1038/sj.bdj.4800776
- 26. Russo PA, Banovic T, Wiese MD, Whyte AF, Smith WB. Systemic allergy to EDTA in local anaesthetic in radiocontrast media. J Allergy Clin Immunol Pract 2014; 2 (2):225–230. doi: 10.1016/j.jaip.2013.12.001
- 27. Sambrook PJ, Smith W, Elijah J, Goss AN. Severe adverse reactions to dental local anaesthetics: systemic reactions. Aust Dent J 2011; 56(2):148-53; quiz 234. doi: 10.1111/j.1834-7 819.2011. 01316.x
- 28. Sampson HA, Muñoz-Furlong A, Campbell RL, Adkinson NF Jr, Bock SA, Branum A, Brown SG, Camargo CA Jr, Cydulka R, Galli SJ, Gidudu J, Gruchalla RS, Harlor AD Jr, Hepner DL, Lewis LM, Lieberman PL, Metcalfe DD, O'Connor R, Muraro A, Rudman A, Schmitt C, Scherrer D, Simons FE, Thomas S, Wood JP, Decker WW. Second symposium on the definition and management of anaphylaxis: Summary report Second National Institute of Allergy and Infectious Disease/Food Allergy and Anaphylaxis Network symposium. J Allergy Clin Immunol 2006; 117(2):391-7. doi: 10.1016/j.jaci.2005.12.1303
- Seymour RA, Meechan JG, Yates MS. Pharmacology and Dental Therapeutics 3rd edition. Oxford: Oxford University Press 1999; 120–124.
- 30. Sheikh A, Ten-Broek V, Brown SG, Simons FE. H1-antihistamines for the treatment of anaphylaxis with and

- without shock. Cochrane Database Syst Rev 2007; 2007(1):C D006160. doi: 10.1002/14651858.CD006160.pub2
- 31. Sheikh A. Gluco-corticosteroids for the treatment and prevention of anaphylaxis. Curr Opin Allergy Clin Immunol 2013; 13(3):263-7. doi: 10.1097/ACI.0b013e32836097f4
- 32. Sicherer SH, Leung DY. Advances in allergic skin disease, anaphylaxis, and hypersensitivity reactions to foods, drugs, and insects in 2012. J Allergy Clin Immunol 2013; 131(1):55-66. doi: 10.1016/j.jaci.2008.12.025
- 33. Simons KJ, Simons FE. Epinephrine and its use in anaphylaxis: Current issues. Curr Opin Allergy Clin Immunol 2010; 10(4):354-61. doi: 10.1097/ACI.0b013e32833b c670
- 34. Thyssen JP, Menne T, Elberling J, Plaschke P, Johansen JD. Hypersensitivity to local anaesthetics update and proposal of evaluation algorithm. Contact Dermatitis 2008; 59(2):69e7 8. doi: 10.1111/j.1600-0536.2008.01366.x
- 35. Vervloet D, Magnan A, Birnbaum J, Pradal M. Allergic emergencies seen in surgical suites. Clin Rev Allergy Immunol 1999; 17(4):459–467. doi: 10.1007/BF02737650
- 36. Wahl MJ, Brown RS. Dentistry's wonder drugs: Local anesthetics and vasoconstrictors. Gen Dent 2010; 58(2):114-2 3; quiz 124-5.
- 37. Webb LM, Lieberman P. Anaphylaxis: A review of 601 cases. Ann Allergy Asthma Immunol 2006; 97(1):39–43. doi: 10.1016/S1081-1206(10)61367-1