Print ISSN: 0003-2778

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Journal of the Anatomical Society of India (ISSN: Print 0003-2778) is peer-reviewed journal. The journal is owned and run by Anatomical Society of India. The journal publishes research articles related to all aspects of Anatomy and allied medical/surgical sciences. Pre-Publication Peer Review and Post-Publication Peer Review Online Manuscript Submission System Selection of articles on the basis of MRS system Eminent academicians across the globe as the Editorial board members Electronic Table of Contents alerts Available in both online and print form. The journal is published quarterly in the months of January, April, July and October.

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Impact Factor[®] as reported in the 2019 Journal Citation Reports[®] (Clarivate Analytics, 2020): 0.227

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Published by

Wolters Kluwer India Pvt. Ltd A-202, 2nd Floor, The Qube, C.T.S. No.1498A/2 Village Marol, Andheri (East), Mumbai - 400 059, India. Phone: 91-22-66491818 Website: www.medknow.com *Printed at* Dhote Offset Tech. P. Ltd Goregaon (E), Mumbai, India

Print ISSN: 0003-2778

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Acceptance of Donor Bodies and Their Embalming During COVID-19 Period: A Challenge to Anatomists

The journey of a medical student begins with learning anatomy by cadaveric dissection. It not only helps to acquire knowledge about the anatomy of a human body but also helps in developing surgical skills and other clinical skills required as a physician.^[1] With the adoption of the Curriculum-Based Medical Education, there has been a change in the teaching methodologies in medical education.^[2] Many medical colleges are now using modern teaching aids such as E-learning, simulation, role play, case study, and virtual dissectors, but nothing can replace the traditional method of teaching by cadaveric dissection.

In the present scenario of COVID-19, when the Ministry of Health and Family Welfare, the Government of India, and the Board of Governors of Medical Council of India have issued guidelines to all the medical institutions to suspend classes for courses such as M. B. B. S., B. Sc., nursing, and other paramedical and allied health sciences,^[3] teaching is now being carried out by taking online lectures and practical demonstrations using video software such as Zoom, Google Classroom, Teams, and YouTube. Even in pre-COVID times, there has been a reduction in number of body donations in most parts of the world; hence, many institutions are adopting alternative models, especially for the teaching of gross anatomy. Hence, conducting anatomy dissection classes online has become very challenging for both teachers and students.

The COVID-19 pandemic has taken a toll on body donations for research in many parts of our country. Medical colleges/hospitals are not accepting donated bodies which have dealt a severe blow to the effort of raising awareness on body donations for scientific research.

Numerous guidelines have been produced by organizations and governments during the present novel coronavirus pandemic, which will be of great use to anatomists who facilitate donor programs.

The COVID-19 virus is mainly transmitted through large respiratory droplets by inhalation or contact with mucosal surfaces, but other modes of transmission have been suggested (airborne, fecal-oral,^[4] and contact with contaminated surfaces).^[5] There is no evidence so far that the virus is transmitted through contact with the skin of a deceased person, but as the virus is known to persist on surfaces for hours or days, depending on the nature of the surface,^[6] this mode of transition cannot be ruled out. The risk of transmission likely increases with direct contact with bodily fluids, and certainly increases with invasive handling of the cadaver, as in autopsy procedures, if it produces droplets or aerosols.^[7] No evidence has been found so far of individuals who have become infected from exposure to the bodies of persons who have died from 2 COVID-19 (WHO 2020b). In general, the potential risk of transmission related to the handling of bodies of deceased persons with suspected or confirmed COVID-19 is considered low^[7] (ECDC 2020a). While there is no evidence yet that the COVID-19 virus is specifically inactivated in a preserved body donor, the commonly used preservatives, formaldehyde and ethanol, appear to be efficient against the virus.^[8]

In general, the safety precautions applied in the basic handling of any human cadaver should cover the risk of a COVID-19 infection. As in any given case, any cadaver should be treated as potentially infectious. In the absence of a test for COVID-19, this also applies to the risk of a COVID-19 infection. All the staff who are responsible for the collection, transportation and preparation of bodies which are infected or suspected of being infected with COVID-19, must be trained specifically for their tasks. They should be well trained in donning and doffing of personal protective equipment (PPE). They should also be trained to use Infection Prevention and Control practices while handling the bodies. The staff should be trained in performing the last rites in a dignified way. The staff must be trained in, and apply, standard precautions for hand hygiene. The staff handling the bodies should take care of proper disinfection of instruments, washing them with detergent, and sterilizing in autoclave. All the instruments should be sterilised chemically with 70% alcohol, diguanide or 2% glutaraldehyde or 1 % hypochlorite. The area of the body receiving unit and of the embalming room and adjacent environment should also be disinfected using chemicals such as 70%-90% ethanol, hypochlorite based products (0.1%), and hydrogen peroxide (0.5%). Any tubes, drains, and catheters on the dead body must be removed with caution; any puncture holes or wounds on the dead body should be disinfected with 1% hypochlorite and dressed with impermeable material; oral and nasal orifices should be plugged. Extreme cautions should be undertaken while using sharp objects, IV catheters, etc., All medical waste should be handled by the following biomedical waste management rules. All surfaces of the isolation area should be wiped with 1% sodium hypochlorite solution giving a contact time of 30 min and then allowed to dry.

Embalming of bodies infected by the novel coronavirus is not recommended by the WHO,^[9] but this is in the context of advice for funeral homes. In the case of anatomy departments, embalming cannot be avoided, especially for cosmetic embalming and for transportation of bodies. The reason provided by the WHO^[9] and the New South Wales Health authority for not recommending embalming is in order to minimize manipulation of the body and thus the possible generation of aerosol. The Department of Health of South Africa (2020) asserts that embalming of a body infected with the novel coronavirus does not pose a risk. However, forced inflation of the lungs, which may occur during fixation, may generate aerosol^[10] (RCP, 2020b). Thus, any aerosol-generating procedures and splashes of contaminated fluids should be avoided during embalming. The use of PPE, as described above, applies during all embalming procedures.

Body donation is voluntary, and there is no compulsion. In this period of COVID-19 pandemic, we can deny acceptance of a donated body even if the body is COVID-19 negative, considering the risk of community transmission via the staff who are handling the body in hospital or the relatives accompanying the body, as there are chances of them being asymptomatic COVID-19 carriers.

Due to lack of clear cut guidelines regarding the acceptance or non-acceptance of a non COVID body, we may accept the donated body, provided the deceased is COVID negative, doesn't come from a containment zone, has no suspected history and is fulfilling other criteria for body donation.

Keeping in view of the risks associated with handling of dead bodies and the necessity of cadaveric dissection on donated bodies, every medical teaching institution should formulate the best practice guidelines for handling body donation and embalming procedures during the novel coronavirus pandemic.

Vishram Singh, Rohini Pakhiddey¹

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Article Info

Received: 05 September 2020 Accepted: 07 September 2020 Available online: ***

Access this article online				
Quick Response Code:	Website: www.jasi.org.in			
	DOI: 10.4103/JASI.JASI_173_20			

How to cite this article: Singh V, Pakhiddey R. Acceptance of donor bodies and their embalming during COVID-19 period: A challenge to anatomists. J Anat Soc India 2020;XX:XX-XX.



Morphometric Variations of Nasal Parameters in Gujarati Population: An Anatomical Study

Abstract

Introduction: Nasal morphology is an important factor in forensic investigations and facial reconstructive procedures. Found to be strongly related to ethnicity and environmental factors, and known to be sexually dimorphic, the study of nasal parameters is useful in forensic facial reconstruction. The aim of the study is to evaluate the nasal morphological characteristics of the Gujarati population as an indicator for personal identification and to assess the prevalent nasal morphology of the study population determined. Material and Methods: The study involved randomly selected 180 healthy subjects (90 males and 90 females) between three age groups. Nasal width and nasal height were measured using a digital vernier caliper, and nasal index was calculated along with other parameters. The data were statistically analyzed. **Results:** The mean nasal width for male and female was 38.23 mm and 34.94 mm while the mean nasal height was 47.59 mm and 44.35 mm, respectively. The mean nasal index for male subjects (81.08) was also higher than for female subjects (77.30). The morphological classification showed the mesorrhine nose type as the most prevalent among both the males (58.88%) and females (66.66%). Discussion and Conclusion: The population under the study exhibits mesorrhine type of the nose and shows sexual dimorphism in the values of nasal measurements. Thus, the current study is valuable not only in forensic facial reconstruction but also as an added method for determining the gender and ethnicity of an unidentified individual.

Keywords: Anthropometry, forensic identification, morphology, nasal height, nasal index, nasal width, sexual dimorphism, variation

Introduction

The qualitative and quantitative assessment of the human body has a long history dating back as far as the middle kingdom where local artisans used square grid patterns to produce standard proportions of human figures establishing different formula for males and females.^[1] Nasal anthropometry involves the systematic measurement and analysis of the human nose which aims to provide baseline data for the study of different human populations.

Nose is a pyramidal structure located in the midline of the mid-face and attached to the facial skeleton made up of bones, cartilages, muscles, and soft tissue.^[2] During racial evolution, nose shows a high degree of variation.^[3] Both genetic and environmental factors influence the shape of the nose.^[4] Due to the natural selection, in cold and dry climates, the nose tends to be narrower, while in warmer and moist environment,

it is broader in humans.^[4] Hence, the anthropometric parameters changes with ethnicity, race, and gender. These baseline data are of great importance in nasal reconstructive surgeries, forensic science, facial reconstruction, and anthropology.

Any alteration in the shape of the nose leads to a gross change in the facial appearance of an individual. Hence, population-specific data regarding the nasal morphology are of utmost importance for the plastic surgeons in reconstructive studies and forensic investigators in facial identification and reconstructions. In any nasal reconstructive procedure, there should be a thorough analysis of the nasal morphology of population of a particular geographic area. Hence, plastic surgeons and forensic investigators both can benefit from a regional database regarding nasal anthropometry.

Nasal index is measured as the ratio of maximum breadth of nose with the maximum length of nose multiplied by 100, which helps characterize nasal morphology

How to cite this article: Rohith MM, Roy J, Johnson A. Morphometric variations of nasal parameters in Gujarati population: An anatomical study. J Anat Soc India 2020;69:XX.

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into five types.^[5] Thus, it is clearly understood that the study and comparison of nasal index form the basis of racial, ethnic, and gender determination among different populations.

It is noted that Berom, Igbos, Yorubas, and Ijaws ethnic tribes from African region fall under the platyrrhine nose type, whereas Western Uttar Pradesh region and South Indian males have mesorrhine type of nose and females have leptorrhine type of nose.^[6-9] At present, there are very little data about the nasal morphology of Gujarati population. Therefore, the present study aims toward measuring and analyzing the nasal parameters of Gujarati population, thereby obtaining the mean nasal index and assessing the sexual dimorphism related to nasal morphology. The study also tries to establish the distribution of basic nose types, shape of the nose, nasal tip angle, and nostril model in Gujarati population to provide the baseline anthropometrical data for nasal reconstructive surgeries, forensic science, and anthropological studies.

Material and Methods

The study group consisted of 180 subjects, comprising 90 males and 90 females, in the age range of 20 years and above who were randomly chosen. They were further divided into three age groups, i.e., 20–40 years, 41–60 years, and 61 years onward [Table 1]. All the subjects were volunteers and were made aware of the study. Written consent was also obtained from each and every individual. The subjects were included in the study after recording demographic data, brief history of the present illness, and medical/surgical history. Apparently healthy individuals with average-to-good built and no visible facial deformity were included, while those with hereditary facial asymmetries, history of esthetic treatment or orthognathic surgery, history of maxillofacial trauma, and history or clinical characteristics of any type



Figure 1: Measurement of nasal width and nasal height

of systemic disorders such as bone diseases, nutritional diseases, and endocrinal diseases were excluded. Total six parameters of the nasal structure were included in this study and they are as follows.

- 1. Nasal breadth
- 2. Nasal height
- 3. Nasal index
- 4. Nose tip angle
- 5. Nostril model
- 6. Nose shape from lateral aspect.

The nasal breadth was measured from the alare (al) of the one side to the alare (al) of the other side by using a vernier caliper. The nasal height was measured from the nasion (n) (a fixed point between the eyes) to the subnasale (nasolabial junction) [Figure 1]. The nose tip angle is an internal angle formed between the line drawn from nasion (n) to pronasale (prn) and the line connecting pronasle (prn) and the subnasale (sn) [Figure 2]. The angle was measured by placing a protractor on the lateral aspect of the nose. The horizontal reference line of the protractor was made parallel to that of the prn–sn line, and the angle formed by the n–prn line in relation to the prn–sn line was recorded [Figure 2]. The nasal index was calculated from the nasal breadth and the nasal height as the formula for nasal index is given below:

Nasal index=
$$\frac{\text{Nasal breadth}}{\text{Nasal height}} \times 100$$

Based on the nasal index, the general nose shape was determined to be one of the following types.^[5]

- 1. Hyperleptorrhine (\leq 54.9) Very long and narrow nose
- 2. Leptorrhine (55-69.9) Long and narrow nose



Figure 2: Measurement of nasal tip angle

	Table 1: A	ge group and nu	mber of partic	ipants		
	1	Number of subject	s		Mean age (years)	
	Male	Female	Total	Male	Female	Total
Group 1 - 21-30 years	30	30	60	23.6	22.07	22.79
Group 2 - 31-40 years	30	30	60	35.78	34.42	35.21
Group 3 - 41 years and above	30	30	60	57.20	51.36	55.37

Journal of the Anatomical Society of India | Volume 69 | Issue 3 | July-September 2020

- 3. Mesorrhine (70–84.9) Moderate nose of moderate size
- 4. Platyrrhine (85–99.9) Broad nose
- 5. Hyperplatyrrhine (≥ 100) Very broad nose [Figure 3].

The nose shape from the lateral aspect and the nostril model were determined by comparing the parameters with the chart provided by a study done by Uzun and Ozdemir^[10] [Figure 4].

Statistical analysis

The data recorded were compiled using MS Excel Spreadsheet (version-2010) and subjected to statistical analysis using Statistical Package of the Social Sciences Version 20. (SPSS 20.0, IBM, Armonk, NY, United States of America). The existence of significant differences between the means of all the parameters for the gender was analyzed using independent Student's *t*-test. One-way ANOVA was also performed to determine any significant changes in the parameter values due to age progression. A P < 0.05 was considered to be statistically significant.

Results

Nasal index

The nasal index showed statistically significant difference between males and females in all the age groups. The nasal index was found to be higher for males than females in all the age groups [Table 2]. Based on the nasal index, the general shape of the nose was also determined. Mesorrhine type of nasal shape was found to be prevalent for both males and females [Table 3].

Nasal width and nasal height

Nasal width and height were found to be significantly different among males and females. Both of the metric parameters were found to be higher for males and lower for females. One-way ANOVA was performed for these parameters which revealed that these parameters were significantly different among the three age groups. The data suggest that these metric parameters increase with age. The results are described in Table 4.

Nasal shape from lateral aspect and nostril model

The nasal shape from the lateral aspect and nostril model was determined by a chart provided by Uzun and Ozdemir.^[10] In case of the nasal shape from lateral aspect, "Type III" was found to be prevalent for males, and "Type I" was common for females [Table 5]. "Type II" nostril model was found to be prevalent for both male and female [Table 6].

Nasal tip angle

The minimum value for this parameter was found to be 45° and the maximum value was 89°. Based on these data, the



Figure 3: Variations in the nasal morphology based on nasal index

Table 2: Nasal index								
Nasal index	Male				Female			
	Group 1	Group 2	Group 3	Total	Group 1	Group 2	Group 3	Total
M	81.89	81.19	80.17	81.08	76.56	78.11	77.25	77.30
SD	9.40	6.98	8.62	8.61	8.45	8.41	8.76	9.02

M: Mean value, SD: Standard deviation

	Tal	ble 3: Prevalence of gen	neral nose shape based o	on nasal index	
	Type I, <i>n</i> (%)	Type II, <i>n</i> (%)	Type III, <i>n</i> (%)	Type IV, <i>n</i> (%)	Type V, <i>n</i> (%)
Male	0 (0)	8 (8.88)	53 (58.88)	27 (30)	2 (2.22)
Female	0 (0)	12 (13.33)	60 (66.66)	16 (17.77)	2 (2.22)

Type I: Hyperleptorrhine, Type II: Leptorrhine, Type III: Mesorrhine, Type IV: Platyrrhine, Type V: Hyperplatyrrhine

Table 4: Mean value and standard deviation of different metric nasal parameters among male and females of different

		age group	55	
Sex	Nasal parameters		Different age groups, mean	±SD
		20-30 years (mm)	31-40 years (mm)	41 years and above (mm)
Male	Nasal width	37.51±2.306	38.44±2.17	38.75±3.10
	Nasal height	46.17±4.335	47.54±3.02	49.06±2.87
Female	Nasal width	33.64±2.54	35.17 ±1.95	36.01±3.30
	Nasal height	43.64±3.24	44.15±3.32	45.28±4.58

SD: Standard deviation



Figure 4: Reference anatomical scale for nasal parameters

Table 5: Prevalence of nasal shape from lateral aspect						
	Type I, <i>n</i> (%)	Type II, <i>n</i> (%)	Type III, <i>n</i> (%)	Type IV, <i>n</i> (%)	Type V, <i>n</i> (%)	
Male	6 (6.66)	13 (14.44)	35 (38.88)	29 (32.22)	7 (7.77)	
Female	45 (50)	9 (10)	3 (3.33)	22 (24.44)	11 (12.22)	

		Table 6: Preva	lence of nostril model		
	Type I, <i>n</i> (%)	Type II, <i>n</i> (%)	Type III, <i>n</i> (%)	Type IV, <i>n</i> (%)	Type V, <i>n</i> (%)
Male	5 (5.55)	55 (61.11)	14 (15.55)	13 (14.44)	3 (3.33)
Female	3 (3.33)	58 (64.44)	17 (18.88)	8 (8.88)	4 (4.44)

nasal tip angle was categorized into five types, and they are as follows:

- Type I − 41°−50°
- Type II $-51^{\circ}-60^{\circ}$
- Type III − 61°−70°
- Type IV 71°–80°
- Type V $81^{\circ}-90^{\circ}$.

"Type IV" $(71^{\circ}-80^{\circ})$ was found to be most prevalent in both males and females [Table 7 and Figure 5].

The data derived from all the parameters can be used to establish the difference in the nasal index among various racial groups. This may help in identification and may be used as an add-on parameter for forensic identification [Table 8].

Discussion

Nose being the most prominent structure of the face is of great significance in both medical science and forensic science. Because of the variations in the nasal parameters, many studies have been carried out on different population to establish the racial and ethnic differences. Previous studies have shown that there is a marked variation of nasal parameters without any specific pattern until 20 years of age. After the age of 20 years, the morphometric changes in the nose occur in a stable manner.^[11,12]

The study conducted on Yoruba tribe of Africa by Oladipo *et al.* suggests that males have mean nasal index of 90 and females have 88.1.^[7] Another study conducted by Esomonu *et al.* shows male and female mean nasal index to be 94.65 and 90.33, respectively.^[13] Patil *et al.* conducted a study on the South Indian population and suggested that the mean nasal index of males was 84.91 and of females was 67.75.^[8] However, the current study on Gujarati populations shows mean nasal index of males to be 81.08 ± 8.61 and mean nasal index of females to be 77.30 ± 9.02.

This study also shows a significant difference in the value of nasal index among males and females, where the Gujarati



Figure 5: Variations in nasal tip angle among Gujarati population

Table 7: Prevalence of nasal tip angle						
	Male, <i>n</i> (%)	Female, <i>n</i> (%)				
Type I - 40°-50°	1 (1.11)	2 (2.22)				
Type II - 51°-60°	7 (7.77)	5 (5.55)				
Type III - 61°-70°	33 (36.66)	28 (31.11)				
Type IV - 71°-80°	39 (43.33)	39 (43.33)				
Type V - 81°-90°	10 (11.11)	16 (17.77)				

Table 8:	Comparison	of	other	studies	with	the	present

	study						
Race/population	Nasal	Author(s), Year					
	index						
Nigerian Igbos	85.7	Akpa et al., 2003					
Ijaws	96.37	Oladipo et al., 2007					
Yorubas	89.2	Oladipo et al., 2007					
Igbos	94.1	Oladipo et al., 2007					
Bheel-Meena	83	Gangrade et al., 2012					
males(Rajasthan)							
Bheel-Meena	79.73	Gangrade et al., 2012					
females(Rajasthan)							
Brahmin Males (Punjab)	70.02	Kaushal et al., 2013					
Brahmin Females (Punjab)	69.89	Kaushal et al., 2013					
Majhabi-Sikh Males	76.51	Kaushal et al., 2013					
(Punjab)							
Majhabi-Sikh Females	68.95	Kaushal et al., 2013					
(Punjab)							
Muslim Males (Punjab)	67.04	Kaushal et al., 2013					
Muslim Females (Punjab)	69.38	Kaushal et al., 2013					
Males(Bekwarra ethnic	94.65	Esomonu et al., 2013					
group)							
Females(Nigeria-Cross	90.33	Esomonu et al., 2013					
River State)							
Hindu males (Gwalior	80.59	Sharma et al., 2014					
region)							
Hindu females (Gwalior	77.29	Sharma <i>et al.</i> , 2014					
region)							
South Indian males	84.91	Patil <i>et al.</i> , 2014					
South Indian females	67.75	Patil <i>et al.</i> , 2014					
Berom males	93.66	Ekwere et al., 2015					
Berom females	92.52	Ekwere et al., 2015					
Males (Western Uttar Pradesh)	75.86	Ray et al., 2016					
Females (Western Uttar Pradesh)	72.08	Ray et al., 2016					
Gujarati males	81.08±8.61	Present study					
Gujarati females	77.30±9.02	Present study					

males were found to be having higher value of nasal index than females. This is in accordance with a previous study conducted by Sharma *et al.*^[5] Another study conducted on the Bheel-Meena tribe by Gangrade and Babel also is in correspondence with this study.^[14] The study conducted by Kaushal *et al.*, on various ethnic groups belonging to Punjab, has also concluded significant sexual dimorphism in the value of nasal index and suggested that males are having higher value for nasal index than females in most of the ethnic groups.^[15]

The present study found the nasal breadth and nasal height of Gujarati males to be higher than that of the females in all the age groups (P < 0.05). This difference of nasal width between males and females was earlier described by Akpa *et al.* (2003) in their study on Igbo tribe.^[16] Esomonu *et al.* also stated this variation in their study on the Bekwarra ethnic group.^[13] A study conducted by Patil *et al.* on South Indian population also corroborates with the result of the current study.^[8]

The present study also concludes that the general shape of nose based on the nasal index was mesorrhine for both males and females. It was also noted that the nasal width and nasal height increase with age. Sforza *et al.* demonstrated similar findings in their skeletal maturity study where they correlated the increase in nasal linear measurements with microscopic changes of the muscle and cartilage of the mid face region.^[17] Numerous studies have also concluded that the elasticity and resilience of the skin due to aging attribute to the increase in the linear dimensions of nose.^[18-20]

From the above data, we can understand that in the Gujarati population, the mean nasal index, nasal height, and nasal width values are having significant sexual dimorphism. Hence, we can use nasal index as a useful parameter for the identification of gender in Gujarati population. Nasal anthropometric and morphologic study of the various ethnic groups of India will help to establish a baseline database of all the nasal parameters as the parameters changes with geographic and environmental conditions. These data will be of great importance in corrective surgeries of the nasal and paranasal structures, forensic facial identification, forensic facial reconstruction, and future anthropological studies.

Conclusion

Nose is the most prominent feature of the face, and it shows significant variation among different races. Thus, it is a reliable parameter for the determination of racial orientation of an individual. The significant differences in the metric and morphologic parameters may also be useful for the medical fraternity for reconstructive or corrective surgery of the nose. Determination of gender is an important aspect of forensic identification which ultimately helps in the establishment of identity of an individual. This study shows significant differences in the mean values of the nasal index, nasal height, and nasal width, which may help in forensic facial identification and gender determination. Thus, it can be stated that the nasal metric and nonmetric parameters are of great importance in medical science, forensic science, and anthropological studies. With further research involving a large sample size, the parameters will be validated. Subsequent studies with sample from various parts of the country will establish the importance of nasal parameters in medical, as well as forensic science.

Declaration of patient consent

The authors certify that they have obtained all appropriate patient consent forms. In the form, the patients have given their consent for their images and other clinical information to be reported in the journal. The patients understand that their names and initials will not be published and due efforts will be made to conceal their identity, but anonymity cannot be guaranteed.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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Original Article



Morphometry of Sphenoid Air Sinus and its Ostium for Surgical Relevance: A Cadaveric Study

Abstract

Introduction: The sphenoid air sinuses and their ostia are highly variable structures. The ostia are located on either side of the midline. The exact location of the sphenoid ostium is very much essential in endoscopic sinus surgeries as well as for the transsphenoidal hypophyseal approach. The present study was done to determine the anatomical location of the sphenoid sinus ostium in relation to the surrounding landmarks. It is a descriptive study. Material and Methods: The study was carried out in the Department of Anatomy, VMKV Medical College and Hospital, Salem, on forty sagittal head and neck sections of adult formalin-fixed cadavers. The length and width of the sinus were measured. The shape and type of the sphenoid sinus ostia were also noted. The distance of the sinus ostium from various landmarks was measured using Vernier calipers. The morphometric parameters were compared by Student's t-test on both sides. Results: Most of the specimens showed sellar variety than presellar variety on both right and left sides. The shape of the sphenoid sinus ostia was predominantly round shape, followed by oval and slit shape. The length and width of the sinus on the right side were less when compared to the left side. The distance between the sphenoidal ostia and several landmarks showed no significant difference except the distance from ostium to superior concha. Discussion and Conclusion: During the endoscopic approach of the sphenoid sinus ostium, it is not always easy to locate the ostium. The present morphometric study will serve as a guideline for the endoscopic surgical approach of the sphenoid sinus ostium using the various anatomical landmarks.

Keywords: Concha, sphenoid ostium, transsphenoidal hypophyseal approach

Introduction

The advancement in endoscopic sinus surgery (ESS) had facilitated a clear cut idea about anatomy and pathophysiology of paranasal air sinuses for surgeons. The surgeon aims to aerate the sinuses and restore the mucociliary clearance so that the paranasal air sinuses are retained back to normal function.^[1] ESS helps the surgeons for the diagnosis and treatment of sinusitis.^[2] The anatomy of sinuses can be visualized clearly by high-resolution computed tomography (CT) which also helps to diagnose the mucosal disease.^[3] The normal anatomy along with any variations of sinus structure, its pathologic appearances in disease condition, has to be well understood by the surgeons and radiologists to provide appropriate patient care.^[4] Anatomic variation of the sphenoid sinus is not so common, but if it is present, then it may complicate the surgery, and henceforth,

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the knowledge of these variations should be known precisely to surgeons. Accurate evaluation of the variations is possible with preoperative CT scans of the sinuses.^[5] A comprehensive knowledge of the variable regional anatomy of the sphenoid sinus will reduce the surgical complications that occur during the transsphenoidal approach and functional ESS.^[6-9] The anatomic variations of the sphenoid sinus predispose to recurrent or chronic sinusitis.^[10]

The body of the sphenoid bone is pneumatized to form the sphenoid air sinus. The pneumatization of bones of the skull base is most common. The sphenoid sinus and adjacent bony structures may show various degrees of pneumatization. Sometimes, a pneumatization can be so extensive that it reaches the anterior clinoid process (ACP), foramen rotundum, vidian canal, pterygoid process (PP), or maxilloethmoid recess. The ACP and PR are the frequent sites of pneumatization in the skull base. The clinical significance of the pneumatization requires clarification.^[5]

How to cite this article: Kumar SB, Selvi PG. Morphometry of sphenoid air sinus and its ostium for surgical relevance: A cadaveric study. J Anat Soc India 2020;69:XX-XX.

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Article Info

Received: 22 August 2019 Accepted: 04 August 2020 Available online: XXXX

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The sphenoid sinus communicates with the nasal cavity through sphenoid ostia, which are located on each side of the midline. The sphenoid air sinus opens into the sphenoethmoidal recess. Sphenoid sinuses and their ostia are highly variable structures. The sinus is related to various structures such as optic nerve in its superolateral wall, carotid artery in the lateral wall, trigeminal nerve in the inferolateral wall, and the vidian nerve in the floor of the sinus.^[11] The above structures are separated from the sinus only by a thin bony wall/septum of around 0.5 mm thickness, and even sometimes, the bony wall is replaced by bony dehiscence.^[12] The pituitary gland lies on the roof of the sphenoidal air sinus which is separated only by a thin bony wall. The sphenoidal air sinus is related to the cavernous sinus and internal carotid artery laterally which may be accidently injured during the transsphenoidal approach of the pituitary gland. The extent of pneumatization of sphenoid air sinus is highly variable.^[13] The exact location of the sphenoid ostium is very much essential in ESSs as well as the transsphenoidal hypophyseal approach. During the endoscopic approach of the sphenoid sinus ostium, it is not always easy to locate the ostium; only few studies were carried out in India on this perspective. The accessibility of the sphenoid sinus is most difficult as it is known for the most variable cavity of the human body.[14] The present study will serve as a guideline for identifying the location of sinus ostium during the endoscopic surgical approach using the various anatomical landmarks.

Material and Methods

The present descriptive study was carried out in the Department of Anatomy, VMKV Medical College and Hospital, Salem, on forty sagittal head and neck sections of adult formalin-fixed cadavers after obtaining ethical clearance from the institution reference number (VMKVMC/IEC/18/51). The length of the sinus was measured from the lower margin of the sphenoid ostium to the midpoint of the posterior wall of the sphenoid sinus, whereas the width was measured from the midpoint of the roof to the midpoint of the floor of the sphenoid sinus. The shape and type of sphenoid sinus ostia were noted [Figures 1 and 2]. The distance of the ostium from the major anatomical landmarks was measured using Vernier calipers. The various surgical relevant anatomical landmarks used include midpoint of choana, nasion, anterior end of the superior concha, and midpoint of basisphenoid [Figure 1]. The extent of pneumatization toward the sella was also noted.

Results

Majority of specimens showed sellar variety (23, 57.5%) than presellar variety (17, 42.5%) on both right and left sides. The shape of the sphenoid sinus ostia was predominantly round shape (16, 40%), followed by oval



Figure 1: Anatomical landmarks for measurements and types of sphenoid air sinus

(12, 30%) and slit shape (12, 30%). The morphometry of sphenoidal air sinus was measured and compared between the right and left sides by independent *t*-test [Table 1]. The length of the sinus on the right side was less when compared to the left side. The width of the sinus showed a significant difference among the sides and was found to be more on the left side when compared to the right side. When analyzed, the morphometric data of the sphenoid air sinus were found to be statistically significant (P < 0.01) [Table 1].

The distance of the ostium from the various anatomical landmarks was measured and compared between the sides, and the distance from the ostium to superior concha was found to be significant, whereas other measurements were insignificant when compared [Table 2]. The various surgical relevant anatomical landmarks used include choana, nasion, anterior end of the superior concha, and basisphenoid.

Discussion

Type of sphenoid sinus pneumatization

The sphenoid pneumatization can be classified as three main types.^[15] Type 1 is the conchal type or the fetal type which shows a small sinus separated from the sella turcica by a very thin layer of trabecular bone. Type 2 is the presellar type or juvenile type and is pneumatized up to the level of sella turcica. Type 3 is the sellar type or adult type and is pneumatized below the sella or further posteriorly.^[10] In a study done by Sareen *et al.*, the sellar type was found to be more frequent. It appears in 75%-86% of cases, followed by the presellar type which is found to be present in 10%-25%, followed by conchal type seen in only 2% of cases.^[16] In a study done by Tan and Ong,^[17] the sellar type was found in 55% of sinuses, followed by conchal (28%) and presellar type (17%). In another study, the sellar pneumatization was most common. Sellar pneumatization is an ideal anatomical configuration for transsphenoidal hypophysectomy because the bulge of the sella turcica floor is easily visualized in the operative field.^[18,19] In the present study, the sellar variety was found in 23 (57.5%), whereas the presellar variety was found in 17 (42.5%) on both right and left sides together, whereas conchal type was not seen in any of the specimens as found by other authors Tan and Ong and Sareen et al.[17,18]

Table 1: Morphometry of sphenoidal air sinus				
Side	Length	Width (cm)		
	(cm)			
Right	1.95±0.23	1.01±0.09		
Left	2.84±0.13	1.68 ± 0.11		
P.Value	0.01**	0.01**		
X 7 1		0 11 00 114 11 100		

Values are expressed as mean \pm SEM, *n*=40 with 20 right side and 20 left side, "Nonsignificant, *Significant, ***P*<0.01. Statistical analysis – Independent *t*-test. SEM: Standard error of mean^[10]

Table 2: Measurement of sphenoidal	ostium	from
various anatomical landmarks		

Side	Mean±SD			
	SO-C (cm)	SO-N (cm)	SO-SC (cm)	SO-BS (cm)
Right	3.30±0.10	4.94±0.27	1.80±0.15	3.94±0.28
Left	3.46±0.22	5.00 ± 0.31	2.43±0.17	4.07±0.21
P.Value	0.72#	0.81#	0.01**	0.75#

SO: Sphenoid ostium, C: Choana, N: Nasion, SC: Anterior end of superior concha, BS: Basisphenoid. Values are expressed as mean±SEM, *n*=40 with 20 right side and 20 left side, #Nonsignificant, **P*<0.05, ***P*<0.01. Statistical analysis – Independent *t*-test^[10]



Figure 2: Type of sphenoid air sinus

Shape of the sphenoid ostia

In a study done by D'Souza *et al.*, the sphenoid sinus ostia were slit type in majority of the cases, followed by oval and round shape.^[10] The round shape ostia were seen more on the left side compared to the right side. In another study, 70% of the ostia were found to be round in shape, whereas 28% were ovoid.^[11] In the study, the sphenoid sinus ostia were predominantly round shape (16, 40%), followed by oval (12, 30%) and slit shape (12, 30%) irrespective of the sides.

Morphometry of sphenoid air sinus

Ariji *et al.* and Ikeda *et al.* reported that the volume of paranasal air sinuses increases up to 20 years and then decreases as age advances.^[20,21] Schatz and Becker observed that sphenoid sinus increases in size up to 15 years and thereafter maintains the same size and

volume.^[22] The sphenoid air sinus starts to develop rapidly after the age of 3 years and spreads back toward the sella turcica around the age of 7 years and reaches adult form at the age of 12 years. The maximum average of sphenoid sinus volume was 7.5 cm³ and 8.2 cm³ reported by Schatz et al.^[23] In another study, the sphenoid sinus volume increased in both sexes up to 25 years $(8.71 \pm 2.44 \text{ cm}^3)$ and decreased after the age of 25.^[2] In the present study, the sphenoidal air sinus was measured and compared between the right and left sides [Table 1]; the volume was not measured as the study was done on specimen of sagittal section and hence only length and width were measured. The length of the sinus on the right side was less when compared to the left side. The width of the sinus showed a significant difference among the sides. When analyzed, the morphometric data of sphenoid air sinus were found to be statistically significant (P < 0.01) [Table 1].

Sphenoid ostium from various anatomical landmarks

The sphenoid sinuses are the most inaccessible paranasal sinuses when compared to all other sinuses.^[24] The distance of the sphenoid ostium from various anatomical landmarks in comparison with other studies is tabulated [Table 3].

In the present study, the distance from ostium to superior concha was found to be significant, whereas other measurements were insignificant when compared between right and left. Kim et al.[25] suggested that the best anatomical reference that enables identification of the sphenoid ostium is the posteroinferior end of the superior turbinate, where each ostium is located medially and superiorly. The sphenoid ostium was identified at an average distance of 1.49 cm superior to the choana by Abuzayed et al.[26] In another study, the distance was 1.37 cm on the right side and 1.5 cm on the left side [Table 3].^[10] In the present study, the distance between sphenoid ostium and choana was 2.30 cm on the right side and 2.46 cm on the left side and was considered to be the most reliable landmark to approach sphenoid ostium. The knowledge of the anatomical relationships of the sphenoid sinus is needed for the success of ESSs and to avoid the various intraoperative surgical complications.^[27]

Conclusion

During the endoscopic approach of the sphenoid sinus ostium, it is not always easy to locate the ostium. The present study will serve as a guideline for the endoscopic surgical approach of sinus ostium using the various anatomical landmarks. The four landmarks include choana, nasion, anterior end of the superior concha, and basisphenoid which will be helping to approach the sphenoid ostium. The study will be further extended in humans using CT scan. Accurate evaluation of the distance of the sphenoid ostium from the above landmarks will be possible with preoperative CT scans of the sinuses.

Table 3: Comparison of distance of sphenoid ostiumfrom various anatomical landmarks with previousstudies

Mean distance	D'Souza et al. ^[10]	Present study
(cm)		
SO-C		
Right	1.37±0.33	2.30±0.10
Left	1.5±0.46	2.46±0.22
SO-SC		
Right	1.52 ± 0.30	1.80 ± 0.15
Left	1.42 ± 0.39	2.43±0.17
SO-N		
Right	4.58 ± 0.48	4.94±0.27
Left	4.49±0.25	5.00±0.31
SO-BS		
Right	3.64±0.63	3.94±0.28
Left	3.62±0.60	4.07±0.21

SO: Sphenoid ostium, C: Choana, N: Nasion, SC: Anterior end of superior concha, BS: Basisphenoid

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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Anomalous Subaortic Left Brachiocephalic Vein: Prevalence and Associated Anomalies

Abstract

Introduction: Thoracic venous anomalies are very rare with an estimated prevalence of 0.7% in the general population. One of the rarest such anomaly is a subaortic course of the left brachiocephalic vein (LBV) commonly associated with congenital cardiac or aortic arch anomalies. Material and Methods: We retrospectively analyzed chest computed tomography scans of 710 participants and found incidentally two cases of anomalous subaortic course of the LBV in two females. Results: The observed prevalence of subaortic LBV is 0.28% (2/710 cases). Isolated subaortic LBV without any associated cardiac or aortic arch anomaly was observed in a female patient. Rare bilateral ectopic origin of bronchial arteries from corresponding subclavian arteries was also noted in this patient. In another female patient, right aortic arch anomaly was associated with subaortic LBV. The right-sided aortic arch had an aberrant retroesophageal left subclavian artery arising from a Kommerell's diverticulum as the last branch. Discussion and Conclusion: Extensive literature search has yielded only 15 cases of isolated anomalous subaortic LBV in subjects without any cardiac and aortic arch anomaly. Although this condition is asymptomatic, its presence, when detected, should alert the clinician to the possible presence of associated congenital cardiac and aortic arch anomalies. Accurate knowledge of these rare anomalies will enhance the diagnostic accuracy and proper interpretation of radiological images. Such thorough knowledge will avoid interpreting the anomalous vein as an enlarged lymph node, enlarged left superior intercostal vein, and reduce the chances of surgical complications.

Keywords: Aberrant left subclavian artery, Kommerell's diverticulum, left innominate vein, right aortic arch, thoracic venous anomalies

Introduction

The left brachiocephalic (left innominate) vein (LBV) is formed by the union of the left internal jugular and subclavian veins at the level of the left sternoclavicular joint. Normally it passes through the prevascular space, from left to right, just above the arch of aorta and anterior to supra-aortic branches to join the right brachiocephalic vein (RBV) to form the superior vena cava (SVC). It is the most anteriorly placed vessel in the superior mediastinum. subaortic Anomalous or retroaortic LBV (ASLBV) is an extremely rare venous anomaly in which the LBV passes below the arch, above the pulmonary artery and behind the ascending aorta to join RBV at or below the level of joining of arch of azygos vein. In such cases, the SVC is short and azygos vein drains into RBV. When present this anomaly is usually associated

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with congenital heart disease (CHD) or aortic arch anomalies like right aortic arch (RAA). Anomalous subaortic LBV is seen in about 0.5%-1.7% of all children with CHD.^[1] It is estimated that about 80% of cases with anomalous subaortic LBV have associated right ventricular outflow tract obstruction.^[2,3] Another common association is the presence of right sided aortic arch.^[4] Isolated subaortic LBV is tremendously rare (0.02%) prevalence) with only a handful of reports available in the literature.^[5] Since this anomaly is asymptomatic, its presence is detected surgery incidentally at autopsy, or radiological investigations. Incidental detection of anomalous LBV should alert the clinician of possible associated CHD or aortic arch anomalies. We report here incidental observation of two cases of anomalous subaortic LBV while retrospectively analysing contrast enhanced computed tomographic (CECT) chest scans.

How to cite this article: Babu CSR, Kumar A, Gupta OP. Anomalous subaortic left brachiocephalic vein: Prevalence and associated anomalies. J Anat Soc India 2020;69:XX-XX.

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Article Info

Received: 17 July 2020 Accepted: 01 September 2020 Available online: ***

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Material and Methods

We retrospectively reviewed CT scans of 710 patients referred to the imaging center for CECT of the thorax for suspected pathologies of the lungs and mediastinum during the period September 2015 to March 2018. The imaging data were collected retrospectively from the archives of the imaging center. The study population included 435 males and 275 females (age ranging from 6 months to 86 years, mean age 51.28 ± 17.70 years). The scans of the patients with malignancies likely to distort the anatomy of aortic arch, poorly enhanced scans, and those with thoracic aortic disease were excluded from the study.

All patients underwent contrast-enhanced multidetector computed tomography by a 64 channel scanner (GE Optima 660, 2011, Tokyo, Japan) and received 90-100 mL of nonionic iohexol contrast (Omnipaque 350 mg I/mL; GE Healthcare, Shanghai, China) at the rate of 5 mL/s intravenously. Written informed consent was obtained by the imaging center from each patient before contrast injection. Sections of 0.625 mm thickness were obtained from the lower part of the neck to the upper part of the abdomen and analyzed in a separate work station (GE: AW Volume share 4.5). After analyzing axial, coronal, and sagittal scans, volume rendered and maximum intensity projections were obtained. The data were collected to study the variations in the branching pattern of arch of the aorta and incidentally two cases having anomalous subaortic LBV were detected.

Results

Two cases of anomalous subaortic LBV were detected incidentally, both in females. Isolated subaortic LBV was detected in a female aged 32 years with a normal left-sided aortic arch. No congenital cardiac anomaly was reported.



Figure 1: Volume rendered image (a) anterior view. Isolated anomalous subaortic left brachiocephalic vein descending on the left side of arch and then below the arch and behind the ascending aorta to join right brachiocephalic vein to form the superior vena cava. (b) Posterior view. The ASLBV passes through aorto-pulmonary window above the right pulmonary artery which is giving an early branch (arrow). Note also the presence of ectopic origin of both left bronchial and right bronchial arteries from the corresponding subclavian arteries (LSA, RSA). AA: Ascending aorta, DA: Descending aorta, PT: Pulmonary trunk

The anomalous vein was crossing the left lateral side of arch and then turning to the right below the arch and above the pulmonary artery [Figure 1a and b]. The anomalous vein coursed anterior to tracheal bifurcation and posterior to ascending aorta to join the RBV just below the insertion of azygos vein [Figures 1b and 2]. An early branch from the right pulmonary artery was also noted [Figure 1b]. Rare ectopic origin of bronchial arteries (BA) bilaterally from the corresponding subclavian arteries was also observed [Figure 1b] Ectopic right BA arose in the root of the neck from the right subclavian (RSA) while ectopic left BA originated from the left subclavian just above the arch [Figure 1b].

The second case of subaortic LBV was seen in a female aged 30 years and was associated with a high positioned right-sided aortic arch [Figures 3 and 4]. The subaortic LBV was passing below the right arch through the subaortic space, above the pulmonary artery and behind the ascending aorta to join the RBV at the level of insertion of azygos vein [Figures 5 and 6]. The anomalous vein was crossing anterior to the trachea [Figure 5c]. Type II RAA with the branching sequence of left common carotid artery, right common carotid artery, RSA, and Aberrant left subclavian artery (ALSA) was identified [Figure 4]. The aberrant left subclavian artery (ALSA) was arising from a retroesophageal Kommerell's diverticulum and was coursing behind the esophagus to reach the left side [Figures 6 and 7]. Descending thoracic aorta was present on the right side of the vertebral column. No cardiac anomaly was seen.

Discussion

Thoracic venous anomalies are very rare with an estimated prevalence of 0.7% in the general population.^[6] ASLBV is



Figure 2: (a-d) Serial axial sections depicting the course of anomalous subaortic left brachiocephalic vein (arrow) passing from the left side of the arch of aorta (a) then behind the ascending aorta and anterior to trachea (b and c) to form superior vena cava (d). DA: Descending aorta



Figure 3: Sagittal section. Higher position of right arch in the superior mediastinum with anomalous subaortic left brachiocephalic vein (blue circle) passing through a larger subaortic space above the pulmonary trunk. AA: Ascending aorta, DA: Descending aorta



Figure 5: (a-d) Serial axial sections showing the course of anomalous subaortic left brachiocephalic vein (arrow) which is sandwiched between ascending aorta and trachea. Note the position of descending aorta on the right side of the vertebra

one such uncommon systemic venous anomaly and when present, is usually associated with CHD, RAA, and other anomalies. It is present in about 0.5%–1.7% of all children with CHD.^[1] The incidence of this venous anomaly associated with CHD was variously reported as 0.2% at necropsy,^[7] 0.5%–0.98% by echocardiography,^[2,8,9] 1.7% by CT,^[4] and 0.57% of surgical cases.^[10] It is estimated that the incidence in the general population is 0.06%–0.37%.^[11] In a review of 1500 angiographic studies of the brachiocephalic veins, Roberts *et al.* reported only one case of subaortic LBV accounting for an incidence of 0.07%.^[12] In a retrospective analysis of CT scans of 81,425 adult patients without CHD, Yamamuro *et al.* estimated an incidence of 0.03%–0.062% for anomalous LBV.^[13] For evaluating the incidence of aberrant LBV and persistent left SVC without



Figure 4: VR image of the right sided arch of aorta with 4 branches – left common carotid artery, right common carotid artery, right subclavian and left subclavian in that order. The left subclavian artery has an aberrant retroesophageal course and exhibits a Kommerell's Diverticulum at its origin



Figure 6: Serial coronal sections showing the course of anomalous subaortic left brachiocephalic vein (arrow). Note the anomalous retroesophageal left subclavian artery is crossing the midline. PT: Pulmonary trunk

bridging vein, Kobayashi *et al.* recently reviewed CT scans of 49,494 patients and found an incidence of 0.055% and 0.15%, respectively, for these two venous anomalies in Japanese patients.^[14] Out of 27 cases of ALBVs detected in their study, only one case had associated CHD, 14 patients had high aortic arches, and in 26 cases, the ALBVs had retroaortic course. Reported prevalence rates were largely derived by the retrospective analysis of the surgical, radiological, and autopsy records [Table 1]. Isolated ASLBV without any associated cardiac and aortic arch anomalies is still rarer,^[5] and some of the adult cases reported since 1980 are summarised in Table 2.

Most commonly, the ASLBV was associated with conotruncal anomalies, aortic arch anomalies and deletion of chromosome 22q11.2. Most common associated finding is tetralogy of Fallot (TOF) reported to be present in 40%–93% cases of ASLBV.^[8-10] Another associated anomaly with high frequency is the presence of right-sided aortic arch (RAA) in 46%–86% of cases.^[2,10] Anomalous subaortic LBV was more common in those with RAA than those with the left aortic arch (P < 0.0001).^[4] The presence of ASLBV in patients with deletion of chromosome 22q11.2 was also reported.^[10,27] Ventricular septal defect (VSD) with

Author year	Modality of study	Cases asso	Cases associated with CHD		Cases without CHD	
		Total number	AS-LBV cases (%)	Total number	AS-LBV cases (%)	
Sherman, 1963 ^[15]	Necropsy	503	2	-	-	
Gerlis and Ho, 1989 ^[7]	Necropsy	2000	3	5000	None	
Choi et al., 1990 ^[8]	Echocardiography	2457	24 (0.98)	-	-	
Curtil et al., 1999 ^[9]	Echocardiography and surgery	5218	27 (0.5)	-	-	
Chen et al., 2005 ^[4]	CT	1812	27	-	-	
Kulkarni et al., 2008[2]	Echocardiography	1432	8 (0.55)	-	-	
Nagashima et al., 2010 ^[10]	Surgical	2449	14 (0.57)	4805	1 (0.02)	
Ko et al., 2011 ^[16]	CT/MRI	-	-	25,940	4 (adult)	
Kahkouee et al., 2017 ^[1]	CT angio	1372	22	-	-	
Yamamuro et al 2017 ^[13]	CT Chest	-	-	81,425	27 (0.033)	
Kobayashi et al., 2018[14]	CT Chest	49,494	1	49,494	26	
Present study	CECT Chest	-	-	710	2 (0.28)	

Table 1: Reported prevalence of aberrant sub-aortic left brachiocephalic vein in cases with and without associated congenital heart disease

CHD: Congenital heart disease, CT: Computed tomography, CECT: Contrast enhanced computed tomography, AS-LBV: Aberrant subaortic left brachiocephalic vein, MRI: Magnetic resonance imaging

 Table 2: Reports of isolated sub-aortic left brachiocephalic Vein in cases of situs solitus (with out any cardiac/aortic arch anomaly)

Author and year	Modality of study	Sex, age	Remarks
Kitamura <i>et al.</i> , 1981 ^[17]	Cadaveric	Male, 69	Left arch
Webb et al., 1982 ^[18]	CECT chest		Left arch
Fujimoto et al., 1992 ^[19]	MRI	Female, 54	Left aortic arch, no cardiac anomaly
Gülsün et al., 2003 ^[20]	CECT, MRA		No cardiac anomaly
Yama et al., 2005 ^[21]	CECT-chest	Male, 46	High left aortic arch
Park et al., 2006 ^[11]	TE Echo, CT	Male, 63	Left aortic arch
Ko et al., 2008 ^[22]	CECT Chest	Male, 74	Normal heart, left arch
		Male, 50	Normal heart, left arch right lung
			cancer
Jayaprakasam et al., 2008 ^[23]	MDCT	Male, 34	No cardiac anomaly
Berko et al., 2009 ^[6]	CT Angio Chest	Male, 50 (1 out of 1000)	Dysphagia, dyspnoea
Nagashima et al., 2010 ^[10]	СТ	Male, 70	Normal heart, left arch distal arch
			aneurysm
Srinivasan et al., 2013 ^[24]	CECT-Chest	Male, 68	Left aortic arch
Hussain et al., 2014 ^[25]	Organ donor		Normal heart
Iimura et al., 2014 ^[26]	Cadaveric	Male, 73	Left aortic arch
Semionov and Kosiuk, 2017 ^[5]	CECT-Thorax	Female, 48	Left aortic arch, normal heart
Kahkouee et al., 2017 ^[1]	CT Angio	Female, 40 (1 out of 1372)	Left arch
Present study	MDCT	Female, 32 (1 out of 710)	Left aortic arch

CT: Computed tomography, CECT: Contrast enhanced computed tomography, MRI: Magnetic resonance imaging, MRA: Magnetic resonance angiography, MDCT: Multi-detector computed tomography

pulmonary atresia, persistent truncus arteriosus, atrial septal defect (ASD), coarctation of aorta, and persistent ductus arteriosus are rare associations observed.^[1,3,10,28,29]

Nakamura *et al.* reported the case of ASLBV in a 62-year-old male with lung cancer and no cardiac anomaly. They emphasized the importance of recognition of this vein in lung cancer patients because misinterpretation as an enlarged superior mediastinal lymph node may cause serious complications.^[30] Botha and Odell reported the case of a 19-day-old male neonate with ASLBV associated with persistence of truncus arteriosus, VSD, secundum

type ASD, absence of arterial duct, and normal left-sided aortic arch.^[28] They noted that Adachi classified ASLBV into two types-type-I anterior to ductus, type II posterior to ductus. Gerlis and Ho added a third type passing below the left pulmonary artery^[7] and Botha's case is Type IV-with the absence of arterial ligament and ASLBV cephalad to the pulmonary artery.^[28] Level of entry of ASLBV into SVC was higher than the insertion of azygos vein in seven cases, at the same level in four cases and lower in three cases out of 14 cases studied.^[31] Hoshino *et al.* described a case of 22q11.2 deletion syndrome with hypoparathyroidism, levocardia, high RAA with aberrant left subclavian artery (ALSA) arising from Kommerell's diverticulum in a 54 year old male.^[27] Hara *et al.* discussed the difficulties encountered while resecting the enlarged lymph nodes



Figure 7: Axial section showing the right aortic arch and the retroesophageal left subclavian artery arising from Kommerell's diverticulum

in a lung cancer patient having ASLBV and RAA with ALSA.^[32] Ohkubo *et al.* reported an anomalous retroaortic LBV with RAA and ALSA in a 49-year-old man.^[33]

Normally, the LBV develops from (a) left anterior cardinal (precardinal) vein and (b) transverse inter-precardinal anastomosis. The exact embryogenesis of the anomalous retro-aortic LBV remains uncertain. Adachi (1933) suggested the presence of double precardinal has anastomoses (ventral superior transverse and dorsal inferior transverse plexuses) around the primitive aorta.^[34] Normally, the ventral superior transverse plexus develops into LBV, but the regression of superior and persistence of the inferior transverse plexus result in the development of anomalous LBV [Figure 8]. Takada et al. thought that this venous anomaly resulted from the precardinal anastomosis being situated posterior to the truncus arteriosus and double (circumaortic) LBV as a consequence of persistence of both cranial ventral and caudal dorsal precardinal anastomosis.^[35] Minami et al. and Kim et al. believed that this anomaly can form secondarily when the elongation



Figure 8: Scheme showing the embryological basis of subaortic left brachiocephalic vein. (a) The developing aorta is surrounded by two venous plexuses connecting right and left anterior cardinal veins. (b) Persistence of ventral superior transverse plexus develops into normal left brachiocephalic vein passing anterior to supraaortic branches. (c) Instead persistence of dorsal inferior transverse plexus and involution of superior plexus gives rise to subaortic left brachiocephalic vein

of the aortic arch prevents the normal cranial ventral precardinal anastomosis to develop.^[36,37] Elongation of the aortic arch leads to narrowing of the prevascular space and widening of the subaortic space (aorticopulmonary window). This widening increases the chances for the persistence of caudal dorsal precardinal anastomosis posterior to the aorta and formation of anomalous LBV.

Arch of aorta and its supraaortic branches develop from paired pharyngeal arch arteries. Normally, the left fourth aortic arch artery develops into the arch of the aorta and right fourth arch artery and part of right dorsal aorta forms part of RSA artery. Distal portion of the right dorsal aorta disappears. RAA is formed due to the involution of left fourth arch artery and persistence of right fourth arch artery and right dorsal aorta. Along with this anomaly if the dorsal inferior transverse venous plexus persists, anomalous subaortic LBV is formed [Figure 9].

In one female patient, we observed anomalous LBV with RAA having aberrant retroesophageal LSA (ALSA) arising



Figure 9: Scheme showing the development of arch of aorta and its branches. (a) Normal development of arch of aorta from left fourth arch artery and left dorsal aorta. (b) Development of right aortic arch due to persistence of entire right dorsal aorta and involution of left dorsal aorta. Persistence of distal part of left dorsal aorta along with left seventh intersegmental artery develops into aberrant left subclavian artery. Anomalous subaortic left brachiccephalic vein develops due to persistence of dorsal inferior transverse plexus

from Kommerell's diverticulum (Type II RAA). Right sided aortic arch is present in 0.05%–0.1% of radiology series and 0.04%–0.1% of autopsy series.^[38] Edwards classified the RAA into three types. Type I is RAA with mirror image branching (left brachiocephalic trunk, right common carotid, and RSA), Type II is RAA with ALSA (left common carotid, right common carotid, RSA, and ALSA) and Type III is RAA with isolation of LSA. Type I and Type III are usually associated with some form of cyanotic CHD like TOF (conotruncal anomalies) but not Type II. Arazińska *et al.* reported that Type II RAA is the most common type of RAA observed in their retrospective analysis (11/20 patients).^[38] Anomalous subaortic LBV combined with RAA and retroesophageal ALSA represented <0.005% in the prevalence of the adult population.

BA originate ectopically from the arch of aorta, subclavian arteries, thyrocervical trunk, and vertebral arteries. Ectopic origin of BA from subclavian arteries bilaterally is very rare.^[39] We have observed bilateral ectopic origin of BA from corresponding subclavian arteries in a female patient.

Conclusion

Although this venous anomaly is asymptomatic having no pathophysiologic repercussions, it should alert the clinician to the possibility of associated CHD and aortic arch anomalies. This variant may mimic an enlarged lymph node resulting in the misinterpretation in imaging studies. This anomaly can cause difficulty during insertion of central venous catheter through a left-sided approach. It can cause difficulties by obstructing the surgical field during corrective surgeries for congenital cardiac anomalies. It can also be mistaken for a persistent left SVC, enlarged left superior intercostal vein or a vertical vein in cases of partial anomalous pulmonary venous return.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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Original Article



The Measurement Indexes and the Relationships with Adjacent Structures of Vidian Canal and Foramen Rotundum using Computed Tomography

Abstract

Introduction: The aim of this study was to evaluate vidian canal (VC) and foramen rotundum (FR) and their anatomical relationships with adjacent structures using computed tomography (CT) in a Turkish subpopulation. Material and Methods: CT images of 150 patients were retrospectively evaluated. Various morphometric measurements (distance from FRs to midline, distance from FR to VC, position and angle of FR, and types of FR and VC) were performed from both left and right sides on CT scans. **Results:** One hundred and fifty patients with a mean age of 41.06 ± 17.812 years were included in this study. The mean distance from midline to right FR was 17.89 ± 1.94 and 18 ± 1.83 in females and males, respectively. The mean distance from midline to left FR was 18.33 ± 1.94 and 19 ± 2.18 in females and males, respectively. Twenty-three cases had Type 1 VC and 40 and 112 cases had Type 2 and 3 VCs, respectively. Three patients had Type I FR, 25 and 57 patients had Type IIa and IIb, respectively, and 93 patients had Type III FR. The position of FRs regarding the base of lateral pterygoid plate was online in 77 patients, medially placed in 92 patients, and laterally placed in 12 patients. Discussion and Conclusion: It is important to know sphenoid sinus and neighboring anatomical structures for planning of endoscopic skull base surgery because it is located close to some important anatomical structures such as internal carotid arteries, optic nerve, and cranial nerves. Surgeons should be careful in preoperative treatment planning and also during the operation.

Keywords: Computed tomography, foramen rotundum, sphenoid sinus, vidian canal

Introduction

The sphenoid sinus has important anatomy during surgical operations of the ventral skull base that is not only the route for reaching the sellar, parasellar, suprasellar, and clival areas but also a way to Meckel's cave and the midcranial fossa. It is located close to various vital anatomical structures such as the internal carotid artery, optic nerve, and cranial nerves. The location of surgical window to the midcranial fossa is in the pterygoid body of the sphenoid bone.^[1]

The foramen rotundum (FR) is a round opening of the sphenoid bone greater wing that anteriorly locates into the pterygopalatine fossa and includes the maxillary nerve;^[2] it locates above and lateral to the pterygoid canal.^[3] The maxillary branch of the trigeminal nerve passes through the FR^[3] after crossing the midcranial fossa where it is in relation to the cavernous sinus lateral wall.^[4] The vidian canal (VC), nerve, and artery are named after Vidius. Vidius was a Medicine Professor at the College of France, and he was a physician to the King of France in the middle of the 16th century.^[5,6] The VC may be observed at the skull base at the anterior border of the foramen lacerum and may locate above and between the pterygoid plates of the sphenoid bone. It enters the pterygopalatine fossa and contains the Vidian nerve, artery, and vein. The Vidian nerve arises from the union of postganglionic sympathetic fibers of the deep petrosal nerve of the carotid plexus and preganglionic parasympathetic fibers from the greater petrosal nerve. After entering the pterygopalatine fossa through the VC, the Vidian nerve enters the posterior area of the pterygopalatine ganglion. The sympathetic fibers provide the vascular constriction in the nasal cavity, and the parasympathetic fibers are responsible for the mucosal secretions in the oral and nasal cavity and the pharynx.^[7,8]

The sphenoid sinus is the most difficult to reach the paranasal sinuses.^[9] In close

How to cite this article: Serindere G, Gunduz K, Avsever H. The measurement indexes and the relationships with adjacent structures of vidian canal and foramen rotundum using computed tomography. J Anat Soc India 2020;69:XX-XX.

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Article Info

Received: 06 April 2020 Accepted: 22 May 2020 Available online: ***

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relationship, there are several vital anatomical structures such as internal carotid artery, optic nerve, and VC. Physicians dealing with skull base surgery must know the important anatomical structures in the skull base. In surgical procedures performed with insufficient anatomical information, complications are inevitable.^[10,11] Imaging modalities are important for the treatment of skull base disorders because clinical evaluation of this region is frequently difficult. Computed tomography (CT) is an ideal choice to evaluate anatomy and is frequently combined with magnetic resonance imaging.^[12]

The aim of this study was to evaluate the measurement indexes of FR and VC and also their relationships with the adjacent structures on CT scans.

Material and Methods

Study design

Before starting the study, ethical approval was obtained from the Local Ethics Committee of Hatay Mustafa Kemal University (Decision date: January 16, 2020, decision number: 07). Images of 150 patients (71 females and 79 males; mean age: 41.06) who applied to the Department of Dentomaxillofacial Radiology were selected from the database. Patients under 18 years of age, or patients with known skull base pathology, remarkable rhinosinusitis (inflammatory changes that prevent the visualization of the skull base anatomy), maxillofacial fractures, sinonasal tumors, or polyposis, were excluded from the study. One hundred and fifty CT images that met the inclusion criteria were evaluated retrospectively.

CT machine (Toshiba Aquilion, Toshiba Medical Systems, Otawara, Japan) was used for all the CT procedures. A routine protocol was performed. All evaluations were performed by a 15.6-inch full HD notebook with resolution of 1920×1080 pixels and by a single observer (GS) who had 6 years. Undecided situations were solved by consensus with KG who had nearly 15 years of clinical experience.

Evaluations were performed using the hospital information system (ENLIL and PACS). For standardization, the first coronal section with VC and FR seen together was selected.

The measurements of this study

An imaginary midline line perpendicular to the rostrum was identified. Various morphometric parameters were studied according to this midline [Figure 1].

These measurements were performed based on the studies of Mohebbi *et al.* as follows:^[13]

- Distance from the midline to the right and left FRs
- Direct distance between VC and FR on both sides
- Horizontal distance between VC and FR on each side (distance between two vertical lines intersecting FR and VC)



Figure 1: The demonstration of various morphometric parameters (red arrow: foramen rotundum, green arrow: vidian canal, and blue arrow: lateral pterygoid plates; V: Vertical distance, H: Horizontal distance, and D: Direct distance) on the coronal computed tomography scan based on the study of Mohebbi *et al.*^[13]

- Vertical distance between VC and FR on each side (distance between two horizontal lines intersecting VC and FR)
- Right and left rotundum angles (were calculated as the angle between the imaginary line connecting FR to VC and the vertical line that crosses VC).

FR's position relative to the base of lateral pterygoid plate as reported by Mohebbi *et al.*^[13] [Figure 2] was as follows:

- Located online: When FR is tangent to the lateral pterygoid plate
- Medially located: If FR is in the medial position relative to the lateral pterygoid plate
- Laterally located: FR is lateral to the lateral pterygoid plate.

The types of FR and VC based on the study of Mohebbi *et al.*^[13] and Lee *et al.*^[10] [Figure 3] were as follows:

- VC was divided into three types on CT:
 - Type 1 VC is completely in the sphenoid sinus
 - Type 2 When the VC is on the floor of the sphenoid sinus or partially protrudes into the sphenoid sinus
 - Type 3 VC fully embedded in the sphenoid corpus.
- FR was divided into three types on CT:
 - Type I FR is completely localized in the sinus cavity
 - Type IIa FR is partially localized in the sinus cavity or partially protruding into the sphenoid sinus
 - Type IIb When FR is tangent to the sinus wall
 - Type III FR is completely inside the sphenoid bone.

Statistical analysis

SPSS version 22 software (SPSS Inc., Chicago, IL, USA) was used to enter and analyze data. All analyses have been done with SPSS version 22 software at an error level of



Figure 2: Coronal computed tomography scan shows foramen rotundum localization with respect to the base of lateral pterygoid plate. (a) Online (right side) and medial (left side) (b) lateral

0.05. The quantitative variables which included distances were expressed as standard deviation and mean. Statistical significance between right and left distances was determined using paired *t*-test and Student's *t*-test. According to the null hypothesis, there was no difference between the groups tested. P < 0.05 was defined as significant.

To study relationship between three qualitative variables of FR-online located, FR-medial, and FR-lateral, the Chi-squared test and Fisher's exact test was used. The relationship between variable of age and all individual quantitative variables was examined with Pearson correlation test. Since the sample size was below 30 at some levels of the qualitative variables and also reached below 5 at some levels of qualitative variables, the age variable distribution was not normal at some levels of qualitative variables with the sample size of below 30. Therefore, the Kruskal-Wallis nonparametric test was used to study the relationship between variable of age and all individual qualitative variables. An Independent *t*-test was used to examine the difference in the variables measuring the right and left sides according to gender. The Chi-squared test was used to examine the relationship between gender and all qualitative variables.

Due to the large sample size and based on the central limit theorem, the average distribution of quantitative variables is approximately normal.

Results

The research participants have a mean age of 41.06 ± 17.812 . Seventy-one (47.3%) females and 79 (52.7%) males were included in this study. Tables 1 and 2 report the descriptive statistics for the quantitative and qualitative variables according to gender, respectively. There was a significant difference in the distance from midline to right and left FR, horizontal, vertical, and direct distances from the right and left FRs to VCs (P < 0.05). There was no difference between the variables FR-online located and FR-medial (P > 0.05). There was a significant difference between the variables Type 1 VC and Type 3 VC. There was a significant relationship between the variables Type 2 VC and Type 3 VC (P < 0.05). There was a significant difference between the variables Type IIb



Figure 3: (1) The types of vidian canal (a) Type 1 (b) Type 2 (c) Type 3 (2) the types of foramen rotundum (a) Type I (b) Type IIa (c) Type IIb (d) Type III

Table 1: Descriptive statistics for the quantitative variables according to gender

Variables	Mean±SD		
	Female	Male	
Age	43.70±15.50	38.68±19.45	
The distance from midline to right FR	17.89±1.94	18±1.83	
The distance from midline to left FR	18.33±1.94	19±2.18	
Direct distance between VK-FR right	6.36 ± 2.42	5.89±2	
Direct distance between VK-FR left	6.54±2.17	6.54±1.75	
Horizontal distance between VK-FR right	4.46±2.01	4.08±1.76	
Horizontal distance between VK-FR left	4.73±1.83	4.75±1.56	
Vertical distance between VK-FR right	4±1.84	3.8±1.54	
Vertical distance between VK-FR left	3.97±1.64	4.05±1.59	
Right rotundum angle	40.66 ± 7.48	37.88±7.16	
Left rotundum angle	40.07±6.75	38.89±6.48	
Direct distance between VK-FR right Direct distance between VK-FR left Horizontal distance between VK-FR right Horizontal distance between VK-FR left Vertical distance between VK-FR right Vertical distance between VK-FR left Right rotundum angle	$6.36\pm2.42 6.54\pm2.17 4.46\pm2.01 4.73\pm1.83 4\pm1.84 3.97\pm1.64 40.66\pm7.48 40.07\pm6.75$	5.89±2 6.54±1.7 4.08±1.7 4.75±1.5 3.8±1.54 4.05±1.5 37.88±7.1 38.89±6.4	

FR: Foramen rotundum, VK: Vitek-Kent, SD: Standard deviation

FR and Type III FR (P < 0.05) [Table 3]. P value between some variables, as shown in Table 3, could not be calculated because the cells of the Chi-squared test were all zero.

Pearson correlation test and Kruskal–Wallis test results are reported in Table 3. There was no significant relationship between age and all individual quantitative variables (P > 0.05).

Table 3 shows that there was a significant difference in age among three levels of left, right, and bilateral localizations of qualitative variables of FR-online, FR-medial, Type 2 VC, Type 3 VC, Type IIb FR, and Type III FR. No significant difference was found in age at levels of other qualitative variables (P > 0.05).

There was a significant difference between gender and the distance from midline to left FR and right rotundum angle (P < 0.05), and no significant difference was found between the other parameters and gender (P > 0.05).

Table 2: Descriptive statistics for the qualitative					
variables according to gender					
Variables	Female, <i>n</i> (%)	Male, <i>n</i> (%)			
FR-online located					
Left	5 (7)	10 (12.7)			
Right	7 (9.9)	6 (7.6)			
Bilateral	22 (31)	27 (34.2)			
Missing	37 (52.1)	36 (45.6)			
FR-medial					
Left	7 (9.9)	4 (5.1)			
Right	4 (5.6)	10 (12.7)			
Bilateral	35 (49.3)	32 (40.5)			
Missing	25 (35.2)	33 (41.8)			
FR-lateral					
Left	1 (1.4)	4 (5.1)			
Right	2 (2.4)	2 (2.5)			
Bilateral	1 (1.4)	2 (2.5)			
Missing	67 (94.4)	71 (89.9)			
Type 1 VC					
Left	2 (2.8)	4 (5.1)			
Right	5 (7)	2 (2.5)			
Bilateral	4 (5.6)	6 (7.6)			
Missing	60 (84.5)	67 (84.8)			
Type 2 VC					
Left	6 (8.5)	3 (3.8)			
Right	4 (5.6)	3 (3.8)			
Bilateral	16 (22.5)	8 (10.1)			
Missing	45 (63.4)	65 (82.3)			
Type 3 VC					
Left	6 (8.5)	5 (6.3)			
Right	5 (7)	5 (6.3)			
Bilateral	37 (52.1)	54 (68.4)			
Missing	23 (32.4)	15 (19)			
Type I FR					
Left	1 (1.4)	1 (1.3)			
Right	0	0			
Bilateral	1 (1.4)	0			
Missing	69 (97.2)	78 (98.7)			
Type IIa FR					
Left	2 (2.8)	0			
Right	5 (7)	7 (8.9)			
Bilateral	6 (8.5)	5 (6.3)			
Missing	58 (81.7)	67 (84.8)			
Type IIb FR					
Left	4 (5.6)	5 (6.3)			
Right	4 (5.6)	4 (5.1)			
Bilateral	15 (21.1)	25 (31.6)			
Missing	48 (67.6)	45 (57)			
Type III FR		~ /			
Left	6 (8.5)	9 (11.4)			
Right	4 (5.6)	4 (5.1)			
Bilateral	36 (50.7)	34 (43)			
Missing	25 (35.2)	32 (40.5)			

FR: Foramen rotundum, VC: Vidian canal

There was no difference between each of the variables and gender (P > 0.05) [Table 3].

Discussion

Although there are some studies on VC in the literature, to our knowledge, no such detailed studies about the radioanatomic relationship of VC and FR with adjacent anatomical structures have been found except one study reported by Mohebbi *et al.*^[13]

Measurement values in our study were lower than those of Mohebbi et al.^[13] It was reported by Mohebbi et al.^[13] that the position of FRs was online in 50% of cases, medially placed in 47%, and laterally placed in 3% of cases. In our study, the case numbers of medially placed FR were higher than online placed, and the incidence of laterally placed FR was higher according to the study of Mohebbi et al.[13] In addition, in their study, it was found that 28 cases (28%) had Type 1 VC and 48% and 24% had Type 2 and 3 VCs, respectively. Four patients (4%) had Type I FR, 28% and 44% had Type IIa and IIb, respectively, and 24% had Type III FR.^[13] In our study, all percentages were found to be lower except the percentage of Type 3 VC and FR. Furthermore, they reported the distance from FR to midline as 19.00 ± 2.07 mm in the right side and 19.34 ± 2.17 mm in the left side. In the right side, horizontal, vertical, and direct distances from FR to VC were reported as 5.89 ± 2.4 mm, 5.06 ± 2.03 mm, and 8.16 ± 2.27 mm, respectively. In the left side, horizontal, vertical, and direct distances from FR to VC were reported as 5.93 ± 2.13 mm, 5.49 ± 2.13 mm, and 9.20 ± 2.15 mm, respectively. Right and left FR angles were reported as 46.76 ± 12.32 and 46.40 ± 10.67 , respectively.^[13] All values of this study were lower than the study of Mohebbi et al.[13]

Kasemsiri *et al.*^[11] reported the average distance from midline to left FR 19.11 mm and to right FR 17.67 mm. Close results were found in this study. Mohebbi *et al.*^[13] reported that the average distance of midline to left FR was found to be significantly more than to right FR. The average horizontal and vertical distances from FR to VC in Kasemsiri's study^[11] showed no significant difference between right and left sides, as the same result in the study of Mohebbi *et al.*^[13] In our study, there was a significant difference in the distance from midline to right and left FR and horizontal, vertical, and direct distances from the right and left FRs to VCs.

Yeğin *et al.*^[14] reported that VC in females was found in 31.5%, 32%, and 36.5% belonging to Types 1, 2, and 3, respectively. VC in males was found in 32.2%, 28.6%, and 39.1% belonging to Types 1, 2, and 3, respectively. In our study, VC in females was found in 15.5%, 36.6%, and 67.6% belonging to Types 1, 2, and 3, respectively. VC in males was found in 15.2%, 17.7%, and 81% belonging to Types 1, 2, and 3, respectively. The reason for some high results in our study may be the calculation of percentages of men and women among themselves, not by total number.

Table 3: Relationships between variables analyzed by different statistical tests stated in the results section of the article

Variables	Р
The distance from midline to right FR	0.000
The distance from midline to left FR	
Direct distance between VK-FR right	0.001
Direct distance between VK-FR left	
Horizontal distance between VK-FR right	0.000
Horizontal distance between VK-FR left	
Vertical distance between VK-FR right	0.304
Vertical distance between VK-FR left	
Right rotundum angle	0.692
Left rotundum angle	
FR-online located and FR-medial	0.000
FR-online located and FR-lateral	0.1
FR-medial and FR-lateral	0.333
Type 1 VC and Type 2 VC	0.250
Type 1 VC and Type 3 VC	0.012
Type 2 VC and Type 3 VC	0.01
Type I FR and Type IIa FR	Not calculated
Type I FR and Type IIb FR	Not calculated
Type I FR and Type III FR	Not calculated
Type IIa FR and Type IIb FR	0.333
Type IIa FR and Type III FR	Not calculated
Type IIb FR and Type III FR	0.001
Age and the distance from midline to right FR	0.395
Age and the distance from midline to left FR	0.803
Age and direct distance between VC-FR right	0.372
Age and direct distance between VC-FR left	0.905
Age and horizontal distance between VC-FR	0.143
right	
Age and horizontal distance between VC-FR left	0.859
Age and vertical distance between VC-FR right	0.410
Age and vertical distance between VC-FR left	0.409
Age and right rotundum angle	0.866
Age and ED online located	0.330
Age and FR-online located	0.023
Age and FR lateral	0.022
Age and Type 1 VC	0.442
Age and Type 2 VC	0.035
Age and Type 3 VC	0.033
Age and Type J FR	0.667
Age and Type IIa FR	0.711
Age and Type IIb FR	0.038
Age and Type III FR	0.049
Gender and the distance from midline to right FR	0.734
Gender and the distance from midline to left FR	0.048
Gender and direct distance between VC-FR right	0.202
Gender and direct distance between VC-FR left	0.984
Gender and horizontal distance between VC-FR	0.217
right	
Gender and horizontal distance between VC-FR	0.934
left	

Table 3: Contd			
Variables	Р		
Gender and vertical distance between VC-FR	0.453		
right			
Gender and vertical distance between VC-FR left	0.738		
Gender and right rotundum angle	0.021		
Gender and left rotundum angle	0.275		
Gender and FR-online located	0.570		
Gender and FR-medial	0.150		
Gender and FR-lateral	0.758		
Gender and Type1 VK	0.376		
Gender and Type2 VK	0.901		
Gender and Type3 VK	0.631		
Gender and Type I FR	1		
Gender and Type IIa FR	0.512		
Gender and Type IIb FR	0.768		
Gender and Type III FR	0.731		

FR: Foramen rotundum, VC: Vidian canal, VK: Vitek-Kent

According to the cone-beam CT study of Bahşi *et al.*,^[15] it was reported that VC totally protruded into the sphenoid sinus (19.75%), partially protruded into the sphenoid sinus (44.37%), and embedded inside bony tissue of the body of sphenoid bone (35.87%). We can say that there are lower results in our study except the incidence of Type III FR.

Yazar *et al.*^[16] reported the mean distance from FR to VC as 7.2 mm. As can be seen, the results of this study were lower than the results in the studies we discussed. We think that this result depends on the subpopulation we are working on.

Nowadays, less invasive endoscopic surgery is frequently performed for treating several diseases of paranasal sinuses.^[14] Surgical excision of pituitary tumors is frequently performed with endoscopic transsphenoidal approach.^[17] The paranasal sinuses have several anatomical variations. Appropriate information of these variations before the surgical operations using CT is fundamental to decrease the complications and to prevent the injury of important structures.^[18] Several vital anatomical structures have a relationship with sphenoid sinus such as VC and FR.^[9]

Conclusion

Although there are studies related to the anatomical and radiological evaluation of VC in the literature, there is no comprehensive radiographic anatomical study in which VC and FR were evaluated together, except for the study of Mohebbi *et al.*^[13] We think that this situation will increase the value of our study, can be a motivation source for other authors who are interested in this subject, and can make useful contributions to the literature.

Financial support and sponsorship

Contd... Nil.

Conflicts of interest

There are no conflicts of interest.

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Prevalence and Pattern of Molar Incisor Hypomineralization in Delhi Region

Abstract

Introduction: Molar incisor hypomineralisation (MIH) is currently the most prevalent of developmental defects of enamel among children. Molar incisal hypomineralisation presents itself as a serious clinical dilemma for pediatric dentists and clinical practitioners. It is a global endemic. However, its prevalence in India remains uncertain to find prevalence and pattern of MIH in Delhi Region. **Material and Methods:** A total of 649 children aged between 7 and 10 years were randomly selected from various schools in Delhi National Capital Region (NCR). The teeth were examined moist under natural light. The developmental defects of enamel were graded using the modified European Academy of Pediatric Dentistry judgment Criteria given by Ghanim *et al.* 2015. **Results:** A total of 97 subjects presented with MIH of 649 subjects examined. MIH presented with a prevalence of 15%. Creamy white opacities were the most common of defects present, followed by yellowish-brown opacities. **Discussion and Conclusion:** The prevalence of MIH in the Delhi NCR region is 15%.

Keywords: Demarcated opacities, developmental defects of enamel, European Academy of Pediatric Dentistry, hypomineralized second primary molar, molar-incisor hypomineralization

Introduction

Molar incisor hypomineralisation (MIH) is defined as the clinical appearance of morphological enamel defects involving the occlusal and/or incisal third of one or more permanent molars and is frequently associated with similar defects in permanent incisors. MIH is as a result of the hypomineralization of systematic origin. The term MIH was first cited by Weerheijm *et al.* 2001.^[1,2]

Any insult or injury to ameloblasts during the formative stage of enamel may lead to qualitative or quantitative defects of enamel. The earlier focus of the research was mainly on developmental defects like amelogenesis imperfect or dental fluorosis, but currently, it has been shifted to MIH due to an increase in the number of cases being reported.^[3]

MIH is a chronological qualitative defect of enamel affecting first permanent molars and being associated with permanent incisors usually presenting as well-demarcated creamy white or yellowish enamel defects, which may or may not be associated with posteruptive breakdown (PEB).^[4] There has been a wide disparity in the prevalence reports of MIH. In national epidemiological surveys on caries prevalence, children are normally not screened for the presence of MIH, so little is known about its occurrence. Various studies done worldwide have shown the prevalence of MIH, ranging from 2.4% to 40.2% due to the different criteria used.^[5]

With the introduction of the terminology of MIH, European Academy of Paediatric Dentistry (EAPD) gave the first index for specific reporting of MIH in 2003. Other Non-European criteria have also reported with wide variations in the prevalence ranging from as high as 40.2% in Brazilian children (Weerheijm *et al.* 2003) to as low as 2.8% in Hongkong (Cho *et al.* 2008).^[6,7]

FDI commission on oral health, research, and epidemiology in 1992 classified enamel defects into two distinct categories: hypomineralized enamel or opacities and-enamel hypoplasia. The first judgment criterion for MIH was given by EAPD in 2003 further modified in 2008.^[8,9]

In 2015 Ghanim *et al.* proposed a charting method for the purpose of using single unified criteria for reporting of MIH cases,

How to cite this article: Singh R, Srivastava B, Gupta N. Prevalence and pattern of molar Incisor hypomineralization in Delhi region. J Anat Soc India 2020;69:XX-XX.

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Article Info

Received: 30 May 2020 Accepted: 31 August 2020 Available online: ***

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which may help preventing the disparity encountered during data compilation due to different coding criteria being used by different researchers, thus provide a better picture of MIH worldwide.^[10]

There have not been any reported studies in India with the new unified criteria to determine the prevalence of MIH; therefore, studies are needed for improved understanding of MIH and its presentation in this region.

The aim of the present study was to determine the prevalence and pattern of MIH in the Delhi Region.

Material and Methods

Study design

Sample size

The present study was conducted on a total of 649 children. Children aged 7–12 years were randomly selected from different schools in the Delhi region.

Inclusion criteria

Normal healthy co-operative children aged 7–12 years, with proper consent duly signed by the parents, were randomly selected for the study.



Figure 1: Normal Incisors



Figure 3: Molar incisor hypomineralisation lesion in incisors

Exclusion criteria

Children having other enamel defects such as amelogenesis imperfecta, dentinogenesis imperfecta, hypoplastic diffused enamel defects, tetracycline stains on index teeth were excluded from the study.

Furthermore, noncooperative children or children affected by any systemic diseases were excluded from the study.

Examination of index teeth

Clinical examination for MIH was conducted in the school itself. Each child was examined by a single examiner under natural light, while seated on a school chair. A trained assistant helped record the findings during intra-oral examination.

The teeth were examined moist. Any debris, if present-were removed with a gauze swab. Four first permanent molar and permanent incisors were examined in each child and evaluated according to base on the modified EAPD charting and recording criteria given by Ghanim *et al.* 2015.

Children with lesions smaller than 2 mm were excluded from further evaluation [Figures 1 and 2]. Teeth which



Figure 2: Normal molar



Figure 4: Molar incisor hypomineralisation lesion in molar

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were currently erupting or erupted $<2/3^{rd}$ were considered to be unerupted. A child was diagnosed positive for MIH if at least one of the erupted first permanent molars presented with a demarcated defect on enamel >2 mm in diameter [Figures 3 and 4].

The data of patients was recorded on a preprinted pro formas with subject's demographic details.

Data were collected manually, tabulated in MS windows excel sheet and analyzed using SPSS software version 17.0 (Inc., Chicago, USA). Data were assessed using Pearson's Chi-square test, Spearman's co-efficient, Fisher's extract, McNemara's test, etc.

Results

A total of 649 subjects were examined, of which 97 subjects (15%) reported with the presence of MIH. No significant differences were found between males and females in MIH prevalence.

Number of subjects with only molars involved were 40 (6.22%), whereas subjects who showed involvement of both molars and incisors affected were 57 (8.78%).

Of the entire subjects involved total number of molars affected were 244/388 (62.89%).

Table 1 shows the distribution of MIH in molars. There was no significant predilection towards right or left side. Mandibular molars were more commonly affected than the maxillary molars.

Of all the teeth involved, a total of 131/776 incisors were affected (16.88%).

Table 2 shows the distribution of MIH in incisors. There was no right or left predilection. Maxillary incisors appeared to be more commonly affected than mandibular incisors.

Table 3 shows the presentation of MIH in molars. Two molars were the most commonly affected followed by the involvement of both maxillary and mandibular molars of both arches simultaneously. Single molar involvement was less common.

Table 4 shows the presentation of MIH in Incisors. Single incisor involvement was the most common, and usually, the maxillary incisors were most involved.

Table 5 shows surface involvement in molars. About 48% of the molars surfaces were affected. Buccal and occlusal surfaces were the most commonly affected. Lingual or palatal surface involvement was less.

- Total number of surfaces-732
- Total number of surfaces affected-352 (48.09%).

Coming to defect characteristics in molars, Creamy white opacities were the most usually encountered in about 81.53% of surfaces involved, followed by yellowish-brown opacities in 18.46%. PEB was also seen in 5.97% of surfaces.

Table 1: Number of molars affected			
	Right	Left	Total
Max	55	52	107
Mand	68	69	137

Table 2: Number of incisors affected					
Tooth	Ri	Right		eft	
	LI	CI	CI	LI	
Max	8	27	31	8	
Mand	8	21	21	7	

Table 3: Number of molar teeth affected as per subject			
	Subjects (n=97)	Molars (%)	
1 molar	14	14 (5.74)	
2 molars	48	96 (39.34)	
3 molars	6	18 (7.38)	
4 molars	29	116 (47.54)	

Table 4: Number of inci	sor teeth affected as per subjects
	Subjects (<i>n</i> =57), <i>n</i> (%)
1 incisor	24 (24.74)
2 incisors	11 (11.34)
3 incisors	13 (13.40)
≥4 incisors	9 (9.27)

Table 5: Number of surfaces affected in molars			
Surfaces	n=352, n (%)	Surfaces	
Occlusal	147 (41.76)	Occlusal	
Buccal	148 (42.05)	Buccal	
Lingual/palatal	57 (16.19)	Lingual	

Discussion

MIH is a term used to describe hypoplastic developmental defects in enamel. The term was first introduced by Weerhejeim *et al.* in 2003. It describes a special pattern of enamel defects affecting the permanent first molars and incisors. The defects present as clearly demarcated opacities ranging from white to yellow to yellowish-brown which may be associated with PEB.

MIH is very frequent in populations across the world. The European academy has long recognized MIH as a global phenomenon necessitating further research and knowledge to have full understanding of the defect. Furthermore, the EAPD had announced criteria specifically for MIH evaluation.

However, there was a wide variation in defect prevalence due to the use of different criteria and indices across the globe; cross comparison of data are difficult. Other non-European countries also showed a wide disparity in the prevalence of MIH, ranging from a low of 2.8% in Hong Kong to a high of 40.2% in Brazil. Considering this, a unified diagnostic criterion was proposed by Ghanim *et al.* in 2015. This criteria proposed a grading method that allowed separate classification for hypomineralized demarcated lesions of the enamel (MIH) and also differentiated it from other similar defects. It also aids in grading the severity of the lesion.

In this study, permanent molars and incisors were scored based on the same criteria. The study reported a prevalence of MIH was 15%, which was in sync with studies done by Kusku *et al.* 14.9%, Laisi *et al.* 16.3%, and Balmer *et al.* 15.9%.^[11,12] It was moderately prevalent than studies done by Bhaskar *et al.* 9.46%, Parikh *et al.* 9.2%.^[13,14] This study showed that MIH in Delhi was moderately prevalent in comparison with data from other cities and countries.

There were differences found between investigations in regard to MIH. This may be because of the reason that the criteria used for investigations were different or self-made at time.^[15] In this study a criterion strictly for MIH was applied and not adapted from other criteria. The modified DDE index does not differentiate clearly between hypomineralisation and certain other enamel defects.

With the standardization of criteria, the prevalence of MIH has shown a gradual increase due to proper diagnosis of lesion and more cases being reported; thus has become a significant concern which needs to be taken care of.

The introduction of EAPD criteria for the evaluation and diagnosis of MIH was considered due to its simplicity which allows easy clinical reproducibility for recording defects. The criterion was standardized by Ghanim *et al.* in 2015, which was used in the present study.

The following study was done in the school campus itself in daylight conditions without using artificial light, which was in accordance with other studies conducted. Given the similarity in age group among different studies, the difference in results may be contributed to differences in criteria used, geographical locations, and other environmental factors.

Of the total subjects affected, about 63% had one or more molars affected. Incisors affected were less commonly affected, about 17% of cases. Although few studies have found a high prevalence of MIH in girls compared to boys, there was no significant gender prevalence in the present study.^[16,17]

Mandibular molars were more commonly affected than maxillary molars. In incisors, there was no significant arch prevalence; it was more or less the same for both arches. The right side appeared to be more commonly effected than the left arch for MIH though the results were not statistically significant.^[18]

Coming to lesion characteristics, white spot lesions were the most common in molars, followed by yellowish-brown lesions. PEB was most commonly associated with yellowish-brown lesions. The results showed a positive association between the change in color of enamel from white to yellowish-brown and the severity of the lesion. This remained constant for all ages. Also with an increase in the number of surfaces involved, there was an increase in the severity of the lesion.

In incisors, central incisors were most affected followed by mandibular central incisors then lateral incisors. The defect was almost 100% confined to the buccal surface and more common in the maxillary arch though not statistically significant. Other studies also reported similar findings where the subjects with an increased no of affected teeth reported with an increase in the severity and extent of lesions Jalevik *et al.* 2001, Leppaneimi *et al.* 2001, Lygidakis *et al.* 2008, Ghanim *et al.* 2011, Parikh *et al.* 2012.^[19-21]

The incisors affected by MIH exhibited demarcated opacities with enamel loss. This was in line with other studies which stated that incisors rarely exhibit postoperative breakdown due to the absence of masticatory forces upon these surfaces.^[22]

In epidemiological studies reporting of the severity of the defect is of greater importance as it reflects upon the treatment needs required by the children affected.

There is a lack of a demarcated classification for MIH lesions, which makes it difficult to diagnose the condition and provide solutions needed.

Conclusion

MIH is the most prevalent current endemic faced by pediatric dentists and clinicians today, which needs to be attended as early as possible.

Further studies are needed in various regions of India to establish the exact current status of MIH in India using the recently standardized modified coding criteria.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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Original Article



Global Research Output of the Cerebellum: Yesterday, Today, and Tomorrow

Abstract

Introduction: The cerebellum has been regarded as a valuable research topic over the years because of its numerous roles in motor and nonmotor functions. The aim of the current study was to provide a comprehensive and holistic evaluation of publications concerning the cerebellum over the last 40 years which can be a potential guide for future research related to the cerebellum. Material and Methods: The Web of Science database was used to identify the publications relating to the cerebellum between 1980 and 2019. The identified publications were analyzed using the bibliometric approaches. The number of the publications concerning the cerebellum to be published between 2020 and 2028 was predicted using the linear regression analysis. Results: The literature review revealed a total of 33,186 publications. The top three active countries were found to be the USA (7362), Japan (2987), and the UK (1994). A positive and significant correlation was found between the number of cerebellum articles produced by the countries and the development indexes of the countries (r = 0.743, P < 0.001; r = 0.676, P < 0.001, r = 0.656, P < 0.001). The top three productive journals were found to be the Journal of Neuroscience, Brain Research, and Cerebellum. The total number of cerebellum publications is expected to continue in an increasing trend and reach a total of 891 publications at the end of 2028. Discussion and Conclusion: Recent studies on the cerebellum have mostly focused on the cerebellar degeneration, functional connections of the cerebellum, and its relationship with various diseases such as ataxia, schizophrenia, Parkinson's disease, autism, and aneurysms.

Keywords: Bibliometric analysis, cerebellum, neuroanatomy, scientometric analysis, trends

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Introduction

The cerebellum, which contains more than 50% of the neurons in the human brain, is a crucial component in the central nervous system. For many years, the cerebellum has been considered as primarily responsible for the coordination of movement and the maintenance of posture and balance. However, recent studies have indicated its contribution to a huge variety of cognitive, emotional, and behavioral functions.^[1-3] Due to its extensive reciprocal connectivity with sensorimotor and association areas of the cerebral cortex, thalamus, and spinal cord, the cerebellum performs various vital functions in humans.^[4] Studies have also shown that structural and functional cerebellar abnormalities are closely related to a wide range of neurodegenerative and psychiatric disorders, including Alzheimer's disease, Schizophrenia, Parkinson's disease, and Multiple sclerosis.[5-8] Due to its vital and different roles in motor and nonmotor functions, the cerebellum has been regarded as a valuable research topic over the past years.

Bibliometrics, based on the application of mathematical and statistical methods, is a commonly used approach for the descriptive analysis of the quality of scientific outputs such as books, articles, and reviews.^[9,10] It provides an objective and effective way for a detailed evaluation of the history, current state and future of scientific publications in a particular field or specific topic, both qualitatively and quantitatively. In bibliometrics, specific features of the scientific documents and publications are analyzed to determine the level of scientific communication. Bibliometric analysis also allows the comparisons to reveal the efficiency of the researchers, institutions, organizations, and countries in various research areas. In this way, bibliometric information enables the identification of global research patterns, dynamics, and trends in a scientific field or topic during a certain period. Therefore, the bibliometric approach reveals valuable and useful information for the comprehensive

How to cite this article: Golpinar M, Demir E. Global research output of the cerebellum: Yesterday, today, and tomorrow. J Anat Soc India 2020;69:XX-XX.

Article Info

Received: 19 June 2020 Accepted: 13 August 2020 Available online: ***

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evaluation of the scientific outputs for both researchers and organizations.^[11,12] There have been many scientific papers concerning the cerebellum over the years, and studies on the cerebellum are still growing rapidly. However, there has been no detailed analysis performed on the scientific output in respect of the cerebellum to date. The aim of the present research was to perform a comprehensive and holistic analysis of the publications concerning the cerebellum published during the period from 1980 to 2019 through bibliometric approaches and to determine the evolving research trends on the cerebellum topic over the 40 years.

Material and Methods

Articles about the cerebellum were accessed through the Web of Science (WoS) database, using the keywords of "cerebellum, cerebellar, cerebelli, and cerebello" in the titles section of the articles (Search codes for repeatability were *Title:* [cerebellum] or *Title:* [cerebellar] or *Title:* [cerebelli] or Title: [cerebello] Refined by: Document Types: [Article] Timespan: 1980–2019. Indexes: SCI-Expanded, SSCI, A and HCI, CPCI-S, CPCI-SSH, BKCI-S, BKCI-SSH, ESCI). All the articles on the cerebellum in WoS downloaded through this search method (accession date: January 15, 2019; the number of publications may vary according to the date of search) were analyzed bibliometrically. A web site (http://lert.co.nz/map/) was used to generate the world map. VOSviewer (Version 1.6.13, Leiden University: Centre for Science and Technology) package program was utilized for bibliometric web visualizations.^[13]

Statistical data analyses were performed using the SPSS software version 22.0 (SPSS Inc., Chicago, IL, USA, license: Hitit University). The normality of the data distribution was tested using the Kolmogorov–Smirnov test. According to the data distribution, the correlation analysis between the number of articles about the cerebellum and the Gross Domestic Products (GDP), GDP (at purchasing power parity) per capita (GDP PPP), and Human Development Index (HDI) were analyzed using the Spearman correlation test. The number of publications to be published in the following years was predicted using the linear regression analysis. Statistical significance was taken as P < 0.05.

Results

The literature review revealed in a total number of 33,186 publications. Of these publications, 22,084 (66.5%) were articles, 6070 (18.291%) were meeting abstracts, 1271 (3.830%) were reviews, 1221 (3.679%) were proceedings papers, 1070 (3.224%) were letters, and the rest were editorial materials (963), notes (824), corrections (181), book chapters (163), early access (38), correction additions (17), discussions (15), news items (13), book reviews (12), reprints (12), books (5), bibliography (1), biographical item (1), hardware review (1), item about an individual (1), and retraction (1). This study performed

the bibliometric analysis of only 22084 publications that were in the article category. Of these articles, 97.2% (21,460) were written in English. Other articles were written in French (184), German (114), Spanish (97), Russian (93), Japanese (51), Portuguese (24), Italian (17), Czech (14), Chinese (9), Turkish (7), Polish (5), Dutch (2), Hungarian (2), Serbian (2), Croatian (1), Slovak (1), and Ukrainian (1).

Development of publications

The annual distribution of the 22,084 articles published between 1980 and 2019 is shown in Figure 1. The figure also displays the regression analysis results indicating the future predictions of expected publication numbers. According to the regression analysis results, it is estimated that 792 (CI 95%: 709–875) articles will be published in the year 2020 and 891 (695–1087) articles in the year 2028 [Figure 1].

Active countries

A total of 142 countries had produced publications. The publication distribution of the countries was displayed on the World Map in Figure 2. The active countries that produced more than 100 articles were the USA (7362), Japan (2987), the UK (1994), Germany (1812), Italy (1771), France (1492), Canada (991), Spain (880), China (854), Netherlands (621), Switzerland (490), Australia (458), South Korea (431), Belgium (361), India (350), Turkey (343), Sweden (319), Brazil (294), Taiwan (274), Denmark (236), Israel (229), Norway (208), Russia (208), Finland (176), Mexico (175), Poland (175), Austria (159), Hungary (135), Greece (112), New Zealand (107), and Iran (100), respectively [Figure 3].

Correlation analysis

A significant positive correlation was found between the number of articles about the cerebellum and GDP, GDP PPP, and HDI development indicators of the countries (r = 0.743, P < 0.001; r = 0.676, P < 0.001, r = 0.656, P < 0.001, respectively).

Active authors

The top ten authors who produced the highest number of articles concerning the cerebellum were as fallow; Timmann D (161), Watanabe M (117), De Zeeuw CI (96), Kano M (94), D'angelo E (91), Hawkes R (90), Schousboe A (75), Mariani J (67), Mikoshiba K (58), and Linden DJ (57), respectively.

Active institutions

The organizations and organizations enhanced that produced the highest number of articles on the cerebellum are listed in Table 1.

Active research areas

In the past 40 years, the top ten research fields of



Figure 1: Number of publications by years on the cerebellum



Figure 2: The distribution of the most productive countries on the cerebellum

cerebellum studies were found to include as follows: Neurosciences (11383; 51.5%), Clinical Neurology (4425; 20.037%), Biochemistry Molecular Biology (1785; 8.083%), Surgery (1410; 6.385%), Pharmacology Pharmacy (991; 4.487%), Physiology (969; 4.388%), Radiology Nuclear Medicine Medical Imaging (915; 4.143%), Cell Biology (899; 4.071%), Multidisciplinary Sciences (890; 4.030%), and Developmental Biology (819; 3.709%), respectively.

Active journals

Articles related to the cerebellum were published in a total of 2327 journals. Of them, 52 were published more than 75 articles [Table 2]. The total number of citations and the mean number of citations per article are also given in Table 2. In addition, the top 10 most active journals that published the articles on cerebellum topic and the top 10 journals with the most citation per article are presented in Figure 4a and b, respectively.

Citation analysis

Table 3 lists the 15 top-cited articles out of the total number of 22,084 articles on the cerebellum. The last column in Table 3 shows the average number of citations per year.

Co-citation analysis

A total number of 343,238 different publications were cited in the references sections of the 22084 articles. The 10 most-cited publications that received more than 400 citations were the studies conducted by Ito, Lowry *et al.*,



Figure 3: The active countries that produced more than 100 articles on the cerebellum



Figure 4: (a) The top 10 most active journals that published articles on the cerebellum. (b) Top ten journals with the most citations per article

Eccles *et al.*, Palay and Chan-Palay, Marr, Schmahmann and Sherman, Albus(1972), Altman, Stoodley and Schmahmann, and Gallo *et al.*, respectively.^[14-23]

Keyword analysis and trend topics

A total number of 25,267 different keywords were used in the 22,084 articles produced about the cerebellum. Table 4 shows at least 53 keywords used in at least 90 different articles. Figure 5 displays the cluster analysis performed with 124 keywords used in at least 50 different articles. Figure 6 demonstrates the network visualization map of the trend keyword analysis in the articles published between 1980 and 2019, the analysis aimed to identify the topic tendency according to the years and potential trends in future. The network visualization map regarding the keyword citation analysis in Figure 7 aimed to identify the effective topics that received more citations.

Discussion

In the present study, we investigated global research output on cerebellum over the last 40 years. The results indicated

Organizations RC Organizations_Enhanced				
Harvard University	314	Centre National De La Recherche Scientifique CNRS	656	
Johns Hopkins University	308	University of California System	645	
CNRS	268	University of London	645	
University Tokyo	268	Institut National De La Sante Et De La Recherche Medicale Inserm	624	
UCL	248	University College London	486	
CNR	226	Harvard University	471	
University Pavia	212	National Institutes of Health (NIH) USA	410	
University Tubingen	205	Sorbonne Universite	390	
Inserm	195	Johns Hopkins University	374	
Kyoto University	191	Assistance Publique Hopitaux Paris APHP	354	
University Minnesota	186	Consiglio Nazionale Delle Ricerche CNR	318	
University California Los Angeles	183	University of Texas System	290	
University Paris 06	176	University of Tokyo	278	
Hokkaido University	173	Erasmus University Rotterdam	259	
University Toronto	172	Riken	243	
Riken	170	Eberhard Karls University of Tubingen	235	
University Texas	167	Sapienza University Rome	229	
University Colorado	166	Max Planck Society	216	
Osaka University	165	University of Toronto	216	
Northwestern University	163	University of Pavia	215	
University California San Francisco	162	Hopital Universitaire Pitie Salpetriere APHP	205	
University Munich	162	Erasmus MC	204	
Washington University	161	University of Munich	203	
Yale University	150	New York University	199	
Erasmus MC	148	Kyoto University	195	



Figure 5: Network visualization map for cluster analysis based on keyword analysis on the cerebellum. Footnote: Six different colors indicate the clustering between the keywords. The size of the circles indicates frequently used keywords. The thickness of the lines indicates that keywords are commonly used together in similar studies

Table 2: Active journals on	the cer	ebellum	
Journals	RC	С	AC
Journal of Neuroscience	704	56,619	80.4
Brain Research	673	21,329	31.7
Cerebellum	640	11,551	18.1
Neuroscience Letters	606	14,139	23.3
Neuroscience	500	18,190	36.4
Journal of Neurochemistry	498	20,305	40.8
Journal of Comparative Neurology	396	19.273	48.7
Journal of Neurophysiology	350	20,095	57.4
European Journal of Neuroscience	315	12.866	40.8
Journal of Physiology London	291	20.452	70.3
Experimental Brain Research	265	10.236	38.6
Developmental Brain Research	257	8282	32.2
Plos One	237	3970	16.8
Journal of Neuroscience Research	237	7047	30.2
Proceedings of the National Academy	233	25 281	111.0
of Sciences of the United States of	220	23,201	111.9
America			
Neuroreport	189	5304	28.1
Journal of The Neurological Sciences	174	3997	23
Neurology	174	11,255	64.7
Neurochemical Research	167	2786	16.7
Neuron	153	19,016	124.3
Acta Neuropathologica	152	4572	30.1
Brain	148	15,216	102.8
Neurosurgery	138	4111	29.8
Neuropharmacology	130	3660	28.2
Journal of Neurosurgery	129	4377	33.9
Journal of Biological Chemistry	126	8872	70.4
American Journal of Neuroradiology	122	4291	35.2
International Journal of Developmental	115	2017	17.5
Neuroscience			
Neurochemistry International	114	1902	16.7
Neuroimage	114	7464	65.5
Journal of Neurology Neurosurgery	106	4305	40.6
and Psychiatry			
Neuroscience Research	106	2011	19
Journal of Neurology	104	1851	17.8
Surgical Neurology	103	1775	17.2
World Neurosurgery	103	394	3.8
Childs Nervous System	102	1278	12.5
Brain Research Bulletin	100	1846	18.5
Experimental Neurology	97	2056	21.2
Molecular Brain Research	96	2865	29.8
Annals of Neurology	94	5975	63.6
Acta Neurochirurgica	03	1504	16.2
Anotomy and Embryology	93	2720	20.2
Anatomy and Emplyology	95	2720	29.5
Communications	90	2322	25.8
Lournal of Dharmanalagu and	00	2600	41
Journal of Pharmacology and	90	3088	41
Experimental incrapeutics	00	1601	107
Seizetice Derest	90	1081	18./
Scientific Reports	88	540	6.2
Alconolism Clinical and Experimental Research	86	2546	29.6
Nescalell			

Table 2: Contd					
Journals	RC	С	AC		
Development	82	8024	97.9		
Journal of Child Neurology	81	1680	20.7		
Journal of Clinical Neuroscience	81	578	7.1		
Molecular and Cellular Neuroscience	81	2421	29.9		
Neurologia Medico Chirurgica	76	527	6.9		

RC: Record count, C: Number of citation, AC: Average citation per document

a general increase in the number of publications concerning the cerebellum in the last 40 years. A total number of 310 articles concerning the cerebellum were published in 1980. The number of articles published per year in the 10-year period was between 310 and 387. The number of articles between 1990 and 1995 was 425-498 per year, and the number of cerebellum-related articles notably increased up to 675 in 1996. The total number of articles, which was approximately 566-688 per year between 1996 and 2012 demonstrated a general increasing trend in recent years and has now reached approximately 800. An analysis of the publication trend of the cerebellum-related articles between 1980 and 2020 showed that although there was a slight decrease in the number of articles between 1998 and 2006, a general increasing trend was noted. Parallel to the growing awareness that the cerebellum is involved in many cognitive and emotional processes as well as a wide variety of neurological and psychiatric conditions, there has been a visible increase in the number of publications in the past decade.^[24-26] In addition, the contribution of researchers from various scientific fields, ranging from cell biology to neurosciences led to increase in the number of publications in this field.^[3,27] When considering similar publication trend from 1989 to 1993, publication trend predicted for the years from 2020 up to 2028 was expected to continue increasing and reach a total of 891 publications per year by the end of 2028. These results indicated that researchers' interest in the cerebellum topic will also continue to increase in the following years.

An analysis of the publication productivity of the countries with respect to the cerebellum showed that the top 10 most productive countries were primarily the USA and Japan, followed by the UK, Germany, Italy, France, Canada, Spain, China, and the Netherlands. Our results showed that the economic size of countries can be directly associated with publication productivity. This hypothesis is confirmed by the highly significant correlation determined between GDP and GDP PPP and the publication productivity of the countries. The significant correlation between HDI and publication productivity was lower than the correlation between GDP and GDP, PPP and publication productivity. This finding which demonstrates that the economic size of the countries was more effective than the development levels in publication productivity. Bibliometric studies conducted in recent years have also shown that

Contd...

No	Article	Author	Journal	PY	ТС	AC
1	Function of the chemokine receptor cxcr4 in haematopoiesis and in cerebellar development	Zou YR et al.	Nature	1998	1886	82
2	Nitric-oxide mediates glutamate-linked enhancement of cgmp levels in the cerebellum	Bredt DS et al.	Proceedings of the National Academy of Sciences of the United States of America	1989	1839	57.5
3	The cerebellar cognitive affective syndrome	Schmahmann JD et al.	Brain	1998	1660	72.2
4	Bell-shaped calcium-response curves of ins (1,4,5) p3-gated and calcium-gated channels from endoplasmic-reticulum of cerebellum	Bezprozvanny I et al.	Nature	1991	1454	48.5
5	Impaired b-lymphopoiesis, myelopoiesis, and derailed cerebellar neuron migration in cxcr4- and sdf-1-deficient mice	Ma Q et al.	Proceedings of the National Academy of Sciences of the United States of America	1998	1275	55.4
6	Marrow stromal cells migrate throughout forebrain and cerebellum, and they differentiate into astrocytes after injection into neonatal mouse brains	Kopen GC <i>et al</i> .	Proceedings of the National Academy of Sciences of the United States of America	1999	1272	57.8
7	Autosomal dominant cerebellar ataxia (sca6) associated with small polyglutamine expansions in the alpha (1a)-voltage-dependent calcium channel	Zhuchenko O et al.	Nature Genetics	1997	1194	49.8
8	Basal ganglia and cerebellar loops: motor and cognitive circuits	Middleton FA et al.	Brain Research Reviews	2000	1175	56
9	Electro-physiological properties of <i>in vitro</i> purkinje-cell dendrites in mammalian cerebellar slices	Llinas R et al.	Journal of Physiology-London	1980	967	23.6
10	Electrophysiological properties of <i>in vitro</i> purkinje-cell somata in mammalian cerebellar slices	Llinas R et al.	Journal of Physiology-London	1980	966	23.6
11	Control of neuronal precursor proliferation in the cerebellum by sonic hedgehog	Wechsler-Reya RJ et al.	Neuron	1999	914	41.5
12	Anatomical evidence for cerebellar and basal ganglia involvement in higher cognitive function	Middleton FA et al.	Science	1994	860	31.8
13	Induction of apoptosis in cerebellar granule neurons by low potassium - inhibition of death by insulin-like growth factor-i and camp	D'mello SR et al.	Proceedings of the National Academy of Sciences of the United States of America	1993	838	29.9
14	Endogenous nitric-oxide release required for long-term synaptic depression in the cerebellum	Shibuki K et al.	Nature	1991	826	27.5
15	International cooperative ataxia rating scale for pharmacological assessment of the cerebellar syndrome	Trouillas P <i>et al</i> .	Journal of the Neurological Sciences	1997	811	33.8

PY: Publication year, TC: Total citation, AC: Average citations per year

economic size is the most effective factor in publication productivity.[28,29]

An evaluation of the journals published articles about the cerebellum showed that the most active journals (i.e., more than 500 articles in total) were the Journal of Neuroscience, Brain Research, Cerebellum, Neuroscience Letters, and Neuroscience. When the total number of citations was considered, the most-cited articles (with more

Table 4: The most used trend keywords on the cerebellu		
Keyword	0	
Cerebellum	5262	
Purkinje cell	620	
Apoptosis	463	

Development	429
Purkinje cells	424
Ataxia	360
Rat	358
Cerebellar granule cells	318
Glutamate	290
Cerebellar ataxia	279
Magnetic resonance imaging	270
Cerebellar cortex	235
Mri	215
Mouse	208
Granule cell	205
Cerebellar granule neurons	185
Cognition	182
Neurotoxicity	182
Gaba	179
Brain	177
Granule cells	176
Posterior inferior cerebellar artery	176
Immunohistochemistry	173
Fmri	165
Oxidative stress	157
Ethanol	153
Calcium	145
Stroke	143
Cerebellar granule cell	136
Motor learning	136
Medulloblastoma	134
Neuroprotection	128
Aneurysm	126
Human	124
Schizophrenia	117
Motor control	115
Aging	114
Cerebellar atrophy	114
Hippocampus	113
Neurodegeneration	107
Nitric oxide	105
Astrocytes	102
Plasticity	102
Cerebellar	98
Cerebellar hemorrhage	97
Cerebellar development	96
Vermis	96
Autism	95
Excitotoxicity	95
Bergmann glia	94
Functional connectivity	94
Purkinje cell	94
Neurons	91

O: Number of occurrences

than 15,000 total citations) were published in the Journal of Neuroscience, Proceedings of the National Academy of Sciences of the United States of America, Brain Research, Journal of Physiology London, Journal of Neurochemistry, Journal of Neurophysiology, Journal of Comparative Neurology, Neuron, Neuroscience and Brain, respectively. On the other hand, an evaluation of the journals published articles about the cerebellum according to the number of citations per article and the citation network map showed that the most active journals were the Neuron, Proceedings of the National Academy of Sciences of the United States of America, Brain, Development, Journal of Neuroscience, Journal of Biological Chemistry, Journal of Physiology London, Neuroimage, Neurology, and Annals of Neurology.

As the above-mentioned journals were the most effective and active journals in terms of the number and quality of the articles produced, these journals could be considered a potential source for the researchers specifically interested in the topic of the cerebellum. These journals could be a good guide for planning new studies or projects about the cerebellum, following current qualified studies, and identifying the most appropriate journal address for the completed studies.

Of the 22,084 articles analyzed, the most-cited study was the article entitled "Function of the chemokine receptor CXCR4 in hematopoiesis and in cerebellar development" written by Zou et al. and published in Nature.^[30] The authors investigated the role of CXCR4, a chemokines receptor, in hematopoiesis and cerebellum development and reported that this receptor had a critical function in the cerebellar neuronal layer formation during the cerebellum development. The second most-cited article was entitled "Nitric-oxide mediates glutamate-linked enhancement of cGMP levels in the cerebellum" written by Bredt and Snyder and published in the Journal of Proceedings of the National Academy of Sciences of the United States of America.^[31] The authors reported that nitric oxide as a messenger molecule mediated the stimulation of cGMP levels in the cerebellum with glutamate.

An analysis of the articles according to the number of citations per year showed that the most-cited study was conducted by Zou *et al.*^[30] Second most-cited article was entitled "The cerebellar cognitive affective syndrome" written by Schmahmann and Sherman and published in the Brain Journal.^[19] The third most-effective cerebellum article according to the average number of citations was the study entitled "Marrow stromal cells migrate throughout forebrain and cerebellum, and they differentiate into astrocytes after injection into neonatal mouse brains" written by Kopen *et al.* and published in the journal of Proceedings of the National Academy of Sciences of the United States of America.^[32] The most-cited articles in the references sections of the 22,084 articles were the studies conducted by Ito, Lowry *et al.*, Eccles *et al.*, Palay and Chan-Palay,



Figure 6: Network visualization map for current trends based on keyword analysis on the cerebellum. Footnote: Indicator shows current publications from blue to red. Keywords in red indicate recently published articles



Figure 7: Network visualization map for the most cited keyword on the cerebellum. Footnote: Keywords in red indicate the most cited keywords

Marr, Schmahmann and Sherman, Albus(1972), Altman, Stoodley and Schmahmann, and Gallo *et al.*, respectively. [14-23]

More qualified and effective studies could be produced if researchers interested in the topic of the cerebellum or planning new studies initially review the above-mentioned studies or the other most-cited papers in Table 3.

According to the keyword cluster analysis findings, cerebellum studies were performed in 6 different clusters which ataxia, Purkinje cell, schizophrenia, development,

aging, and apoptosis in cluster centers. The ataxia, Purkinje cell, and apoptosis centered clusters were predominant as the major research clusters.

An evaluation of the analysis results with a view to identifying the trend topics about the cerebellum articles from past to present showed that the most frequently used keywords around 1990 were magnetic resonance imaging (MRI), stroke, granule cell, hippocampus, plasticity medulloblastoma, neurogenesis, cerebellar atrophy, and electron microscopy. The most frequently used keywords around 1995 included rat, oxidative stress and cerebellar ataxia; and the keywords around 2000 were cerebellar granule cell (cells), glutamate, cognition, immunocytochemistry, and in situ hybridization, while the most frequently used keywords around 2005 included motor learning, motor control, apoptosis, aging, neurotoxicity, neuroprotection, GABI, development, mouse, cerebellar cortex, and calcium. The most frequently encountered keywords during the period from 2010 to 2019 included ataxia, posterior inferior cerebellar artery, aneurysm, MRI, functional MRI (fMRI), diffusion tensor imaging, transcranial magnetic stimulation, functional connectivity, cerebellar degeneration, schizophrenia, Parkinson's disease, autism, neurodegeneration, and multiple system atrophy. An analysis of the keywords used in cerebellum studies conducted after 2010 indicated that there was a trend in the cerebellar functional connectivity, cerebellar degeneration, and relationship of the cerebellum with various diseases and the use of the neuroimaging techniques.

When the citation analysis results of the cerebellum articles were evaluated, the most-cited keywords from past to present included motor control, timing, cognition, language, basal ganglia, autism, mouse, synapse, immunocytochemistry, mitochondria, brain-derived neurotrophic factor, N-methyl-D-aspartate, proliferation, cerebral cortex, granule cell, and astrocyte. Since these keywords were in the most-cited studies with a high impact level, it would be beneficial to consider these in planning research on the cerebellum. An analysis of the keywords used in the cerebellum articles to date showed that the cerebellum was included in several disciplines and studies that demonstrated different approaches in terms of topic and content.

As the WoS database indexes only the journals in the Science Citation Index Expanded and Emerging Science Citation Index, it is generally regarded as having more reliability than other databases such as PubMed and Scopus.^[10,28] Therefore, this study did not use databases such as PubMed and Scopus, but only utilized the WoS database.

Conclusion

The present study is the first attempt to provide a comprehensive analysis of scientific publications regarding

the cerebellum since 1980. The study identified the most active countries, collaborations between countries, the journals with the highest number of and most qualified articles in the cerebellum-related publications, the most-cited cerebellum studies, and the most frequently used keywords by years. Recent studies concerning the cerebellum have mostly focused on the cerebellar degeneration, functional connections of the cerebellum and its relationship with various diseases, primarily ataxia, schizophrenia, Parkinson's disease, and autism, aneurysms and the use of fMRI, MRI, and diffusion tensor monitoring. The comprehensive analysis and the findings obtained in the present study, which covered a long period, can be considered to be of value as a good and useful guide for researchers interested in this topic.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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A Study of Microscopic Changes in the Placenta in Gestational Diabetes Mellitus

Abstract

Introduction: Gestational diabetes mellitus is a complication connected with pregnancy defined as any degree of glucose intolerance that appears during pregnancy with normal values before and usually after the pregnancy. We investigated histological changes of the gestational diabetic placenta and non-diabetic placenta. Research work accomplished in the department of Anatomy, government medical college and superfacility hospital Azamgarh. Placenta collected from the labour room/operation theatre of the department of Obstetrics and Gynecology, government medical college and superficiality hospital, Azamgarh. Material and Methods: Sixty-two freshly delivered placentas were collected - 31 placentas from diabetic mother and 31 placentas from non-diabetic mother (control group). All parturient were aged between 20 to 44 years. This placenta measured on a weighing machine graduated in grams. An approximately 5mm piece of the diabetic placenta was taken and processed for histological examination. Results: Microscopic examination explained dilated blood vessels, subtrophoblastic basement membrane thickness and chorangiosis present in all gestational diabetic placenta. Vessels thrombosis present in 83.87 per cent of the gestational diabetic placenta. Nucleated RBCs. were present in 93.54 per cent of the gestational diabetic placenta. Discussion and Conclusion: Gestational diabetes mellitus induces significant changes in the placenta, both gross and histologic. Effective glycemic regulation is better options for reducing anomalies that cause gestational diabetes. We find dilated blood vessels and necrosis in 100 percent of gestational diabetic placenta and 9.60 percent of nondiabetic placenta. The thickness of the subtrophoblastic basement membrane was present in 100% of the gestational diabetic placenta and 16.12% of the non-diabetic placenta. Vessel thrombosis occurs in 83.87% of gestational diabetic placenta, and 12.90% of non-diabetic placenta. In 93.54 percent of gestational diabetic placenta and 32.25 percent of non-diabetic placenta, nucleated RBC occurs.

Keywords: Chorangiosis, gestational diabetes mellitus, microscopic, placenta

Introduction

The placenta is a fetomaternal organ. It consists of two components: (a) fetal component and (b) maternal component. Fetal component develops from chorion frondosum and maternal component develops from decidua basalis. This is a complex organ of short life span. This is responsible for the transfer of nutrients and waste products between fetal and maternal circulations. The placenta must integrate signals from fetus and mother in an effort to match fetal demand with maternal nutrient.^[1,2] It actually plays an important role in fetal growth and development. Maternal disease that results in abnormal growth and development of the placenta during early pregnancy and mid-pregnancy

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is directly associated with decreased fetal growth and development in the late pregnancy.^[3] Gestational diabetes mellitus (GDM) is a complication connected with pregnancy defined as any degree of glucose intolerance that appears during pregnancy with normal values before and usually after the pregnancy. Diabetes mellitus in pregnancy is known as GDM because woman is nondiabetic before pregnancy. glucose intolerance Any degree of commencement during pregnancy is known as GDM.^[4,5] GDM represents nearly 90% of all pregnancies complicated by diabetes mellitus.^[6] It affects 2%-5% of all pregnancies.^[7] GDM during pregnancy produces varieties of placental abnormalities such as distension of basal membrane of trophoblast proliferation of cells of the endothelium and decreased vascular surfaces of terminal villi. These changes depend

How to cite this article: Singh V, Ranjan K, Tewarson SL. A study of microscopic changes in the placenta in gestational diabetes mellitus. J Anat Soc India 2020;69:XX-XX.

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Article Info

Received: 13 July 2020 Accepted: 02 September 2020 Available online: ***

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Figure 1: Dilated blood vessel and necrosis present in 100% of the gestational diabetic placenta and 9.60% of the nondiabetic placenta



Figure 3: Vessels thrombosis present in 83.87% of the gestational diabetic placenta and 12.90% of the non-diabetic placenta



Figure 5: Chorangiosis present in 100% of the gestational diabetic placenta and 3.22% of the non- diabetic placenta

upon the quality of glycemic control achieved during the critical period of the placental development.^[1] An alteration of the placental function due to uncontrolled GDM results in disturbances of the growth and development of fetus macrosomia, congenital malformations, and intrauterine growth retardation.^[8,9]

Material and Methods

Sixty-two freshly delivered placentas were collected -31 placentas from the diabetic mother and 31 placentas from the



Figure 2: Subtrophoblastic basement membrane thickness present in 100% of the gestational diabetic placenta & 16.12% of the nondiabetic placenta



Figure 4: Nucleated RBCs present in 93.54% of the gestational diabetic placenta and 32.25% of the non-diabetic placenta.



Figure 6: (a) Fetal surface of the gestational diabetic placenta. (b) Maternal surface of the gestational diabetic placenta

nondiabetic mother (control group). All parturient were aged between 20 and 44 years. Research work was accomplished in the Department of Anatomy, Government Medical College and Superfacility Hospital, Azamgarh. Placenta was collected from the labor room/operation theater of the Department of Obstetrics and Gynaecology, Government Medical College and Superfaciality Hospital, Azamgarh. There were no differences according to race culture or environment. The placenta was obtained either from the vaginal route or cesarean section. After washed with the running tap water and dried with the blotting paper, the collected placenta was weighed from the weighing machine graduated in the grams. We took morphometric and macroscopic examination; after that, the placenta was kept



Figure 7: (a) Fetal surface of the nondiabetic placenta. (b) Maternal surface of the nondiabetic placenta



Figure 9: (a) The histological slide of the gestational diabetic placenta - Arrow showing subtrophoblastic basement membrane thickness, ×40. (b) The histological slide of the nondiabetic placenta - Arrow showing basement membrane of the trophoblast, ×40



Figure 11: (a) The histological slide of the gestational diabetic placenta - Arrow showing nucleated red blood cell, ×40. (b) The histological slide of the nondiabetic placenta - Arrow showing nonnucleated red blood cell, ×40

in 10% formalin for 24 h for fixation. After 24 h, this fixed tissue of the placenta passes through a series of procedure from dehydration and clearing to wax impregnation before being sectioned from the microtome. Time took for processing was 24 h, and using microtome instrument, tissues were sectioned at 5 μ m. APES-coated glass slide was used for sectioned tissues. The slide was placed on the hot metallic plate at 60° for 30 min before staining. After that, hematoxylin and eosin staining was done. Slides were prepared and they are passed through xylene-1 for 10 min and xylene-2 for the next 10 min. After that, slides were placed into 100%, 95%, 80%, and 70% alcohol for 5 min each to dehydrate the slides. Slides were placed into the Harris hematoxylin solution for 5–10 min and rehydrate with water thereafter. Again, tissues were washed



Figure 8: (a) The histological slide of the gestational diabetic placenta - Arrow showing dilated blood vessel and necrosis, ×100. (b) The histological slide of the nondiabetic placenta - Arrow showing blood vessels, ×40



Figure 10: (a) The histological slide of the gestational diabetic placenta - Arrow showing vessel thrombosis, ×40. (b) The histological slide of the nondiabetic placenta - Arrow showing blood vessel without thrombosis, ×40



Figure 12: (a) The histological slide of the gestational diabetic placenta - Arrow showing chorangiosis in the chorionic villi, ×40. (b) The histological slide of the nondiabetic placenta - Arrow showing fetal blood vessels in chorionic villi, ×40

with tap water and then placed into 1% eosin for approximately 1-2 min. Finally, tissues were dehydrated by passing through increasing concentration of alcohol and cleared in the solution of xylene for 5 min. Prepared slides were mounted in DPX and covered with the cover slip. Through binocular light microscope, slides were examined at $\times 10$, $\times 40$, and $\times 100$.

Results

Dilated blood vessel and necrosis present in 100% of the gestational diabetic placenta and 9.60% of the nondiabetic placenta shows in Table 1.

Table 1: Presents a comparison of the diabetic (gestational diabetes) and non-diabetic placenta histological changes

histological changes				
	Nondiabetic (control) (%)	Diabetic (gestational diabetes mellitus) (%)		
Dilated blood vessels				
and necrosis				
Present	9.60	100		
Absent	90.30	0		
Subtrophoblastic				
basement membrane				
thickness				
Present	16.12	100		
Absent	83.87	0		
Vessels thrombosis				
Present	12.90	83.87		
Absent	87.09	16.12		
Nucleated red blood cell				
Present	32.25	93.54		
Absent	67.74	6.45		
Chorangiosis				
Present	3.22	100		
Absent	96.77	0		

Subtrophoblastic basement membrane thickness presents in 100% of the gestational diabetic placenta and 16.12% of the nondiabetic placenta.

Vessels thrombosis presents in 83.87% of the gestational diabetic placenta and 12.90% of the nondiabetic placenta shows in Table 1.

Nucleated red blood cell (RBCs) present in 93.54% of the gestational diabetic placenta and 32.25% of the nondiabetic placenta shows in Table 1.

Chorangiosis presents in 100% of the gestational diabetic placenta and 3.22% of the nondiabetic placenta shows in Table 1.

Hence, in our study, chorangiosis subtrophoblastic basement membrane thickness and dilated blood vessels and necrosis present in 100% of the gestational diabetic placenta [Table 1].

Discussion

GDM is a complication connected with pregnancy defined as any degree of glucose intolerance that appears during pregnancy with normal values before and usually after the pregnancy. Diabetes mellitus in pregnancy is known as GDM because woman is nondiabetic before pregnancy. Any degree of glucose intolerance commencement during pregnancy is known as GDM.^[4,5] The disturbance in the adaptation in the metabolism of carbohydrate during the pregnancy causes GDM. The poorly controlled gestational diabetic placenta shows villous edema and marked fibrin thrombi in the syncytiotrophoblast. Brudenell and Doddridge^[10] suggested that villous edema is common in the diabetic placenta. Shen-Schwarz et al.[11] observed similar findings. The diabetes mellitus influences organ system such as cardiovascular excretory and central nervous system.^[12] Diabetes mellitus during pregnancy affects both mother and fetuses.^[13] When metabolic control is good, the mortality rate reduces in the normal population.^[14] The macrosomia and large babies are common in GDM.^[15] The villous immaturity are most common in the GDM.[16] We observed that the gestational diabetic placenta shows dilated blood vessel and necrosis in 100% of cases. However, the nondiabetic placenta shows dilated blood vessel and necrosis only in 9.60% of the placenta. According to the study of Jauniaux and Burton (2006),^[17] GDM causes dilatation of blood vessels and necrosis of the blood vessels. Subtrophoblastic basement membrane thickness increases in the placenta due to GDM. We observed that 100% of the gestational diabetic placenta has subtrophoblastic basement membrane thickness and 16.12% of the nondiabetic placenta has subtrophoblastic basement membrane thickness [Table 1]. According to Hirota^[18] and Okudaira et al.,^[19] the poorly controlled GDM thickness of the basement membrane of syncytiotrophoblast increases. The basement membrane thickness in the gestational diabetic placenta is due to the deposition of mucopolysaccharide. We observed that thrombosis presents 83.87% of the gestational diabetic placenta and 12.90% of the nondiabetic placenta shows in Table 1. We observed that 93.54% of the gestational diabetic placenta has nucleated RBC and 32.25% of the nondiabetic placenta has nucleated RBC [Table 1]. According to Daskalakis et al.,[20] nucleated RBC presents more in the gestational diabetic placenta. We observed that chorangiosis presents in 100% of the gestational diabetic placenta and 3.22% of the nondiabetic placenta [Table 1]. Verma et al.[21] observed chorangiosis seen more in the gestational diabetic placenta as compared to the control. The GDM causes reduced flow of blood through the intervillous space. This reduced supply of the blood is due to vascular compromise.^[2] The angiogenesis occurs due to uncontrolled GDM. 25% surfaces of the villi were taken up by the capillary bed in the gestational diabetic placenta and 50% surfaces of the villi were taken up by the capillary bed in the nondiabetic placenta. Maternal hyperglycemia may impact on the vascular permeability of the placenta. Hyperglycemia has shown to have a direct effect, acting as a proconstrictor,^[22] procoagulator,^[23] proinflammatory,^[24] proangiogenic^[25] and propermeability agent.^[26] According to Lal et al.^[27] the gestational diabetic placenta showed degenerative changes (dilated blood vessels and necrosis), vessel thrombosis, subtrophoblastic basement membrane thickness, and presence of nucleated RBC. This observation supports our findings. According to Verma et al.,^[28] finding of the histological examination such as fibrinoid necrosis, villous edema, villous fibrosis, and chorangiosis more often present in the gestational diabetic placenta compared with the control. This finding supports our findings. Hence, in our study, chorangiosis, subtrophoblastic basement membrane

thickness, and dilated blood vessels and necrosis present in 100% of the gestational diabetic placenta [Table 1].

Conclusion

Diabetes mellitus causes profound gross as well as histological changes in the placenta. The good glycemic control is better options for reducing diabetes-induced abnormalities. In our research work, we found dilated blood vessel and necrosis in 100% of the gestational diabetic placenta and 9.60% of the nondiabetic placenta [Table 1]. Subtrophoblastic basement membrane thickness present in 100% of the gestational diabetic placenta and 16.12% of the nondiabetic placenta [Table 1]. Vessels thrombosis presents in 83.87% of the gestational diabetic placenta and 12.90% of the nondiabetic placenta [Table 1]. Nucleated RBC presents in 93.54% of the gestational diabetic placenta and 32.25% of the nondiabetic basement membrane thickness, and dilated blood vessels and necrosis present in 100% of the gestational diabetic placenta.

Ethical approval

The study was approved by the institutional ethics committee.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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Evaluation of Vascular and Neural Anatomy of the Hand in Adult Cadavers

Abstract

Introduction: We aimed to investigate the vascular and neural anatomy of the hand in adult cadavers and to determine the normal anatomical structure and variations of these structures. Material and Methods: Twenty hands of adult cadavers were examined by the dissection method. After the superficial palmar arch (SPA), the ulnar nerve (UN), median nerve (MN), and deep palmar arch (DPA) were identified, their anatomical structures and variations were evaluated. Then, they were classified according to the number of branches they give off and the course of these branches. Results: When the number of the common palmar digital arteries emerging from the SPA was examined, four branches were observed to emerge from the arch in 80%, and five branches in 20%. When the innervation of the MN and UN was evaluated, 3.5 + 1.5 fingers innervation was observed in 90%, 3 + 1.5 fingers innervation and 3 + 2 fingers innervation were observed in 5% of the samples. When the DPA samples were examined, four metacarpal palmar arteries were observed to emerge from the arch in all of the samples. According to the number of branches they gave off and the course of these branches, the SPA was gathered under seven groups, the nerves were gathered under four groups, and the DPA was gathered under three groups. Discussion and Conclusion: We believe that the data obtained in our study will be used, especially in anatomy education and will guide neurologists, surgeons, orthopedists, radiologists, and anatomists in their studies, diagnosis, and treatments.

Keywords: Classification, deep palmar arch, median nerve, superficial palmar arch, ulnar nerve

Introduction

The hand plays an important role in the daily life of humans. The hand enables us to position, orient, and grasp objects by touching them and provides our communication with the outside world.^[11] In addition to the innervation of the hand and hand-related structures, which take such an essential place in our lives, its nutrition is also vital.

The superficial palmar arch (SPA) and deep palmar arch (DPA), which are formed by the ulnar artery (UA) and radial artery (RA), are the structures feeding the hand.^[2] Most of the studies on the SPA have been performed by the anatomical dissection method.^[3-5] Furthermore, there are also studies conducted with angiography^[6,7] and Doppler-ultrasonography^[8] methods. In our literature review, few studies on the DPA were encountered.^[3,4,9] In the Turkish population, there are few studies using the

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anatomical dissection method concerning the SPA^[10,11] and DPA.^[12]

The extrinsic and intrinsic muscles of the hand provide hand movements. The extrinsic muscles of the hand are innervated by the radial nerve (RN), ulnar nerve (UN), and median nerve (MN). The intrinsic muscles of the hand are innervated by the UN and MN.^[2] There are many studies on the innervation of the extrinsic muscles^[13-15] and intrinsic muscles^[16-19] of the hand. When the studies on the intrinsic muscles of the hand were examined, it was seen that their dates back to very old times and these studies about the motor innervation of muscles. In our literature review, it was seen that there a few studies about the cutaneous distribution of nerves in the palm.[20-22] However, no studies about typing these nerves according to the distribution of fingers were found.

This study's goal was to investigate the normal anatomy and variations of the vascular and neural structures of the hand by examining the hands of adult cadavers by the anatomical dissection method.

How to cite this article: Kastamoni Y, Anil A, Peker T, Anil F. Evaluation of vascular and neural anatomy of the hand in adult cadavers. J Anat Soc India 2020;69:XX-XX.

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Article Info

Received: 07 November 2019 Revised: 13 August 2020 Accepted: 19 August 2020 Available online: ***

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Material and Methods

Twenty adult cadaver hands, which had been previously fixed in a 10% formaldehyde solution, were examined by the anatomical dissection method. Approval was obtained from the Clinical Research Ethics Committee for this study (Date: October 26, 2015, Decision No: 40).

Dissections were performed under a surgical microscope (Carl Zeiss Opmi Pico, Germany) at \times 5 magnification. To remove the skin of each hand, two transverse incisions passing through the wrist fold and finger roots and a vertical incision joining the midpoint of these two incisions were made. After the skin was removed, subcutaneous adipose tissue was cleaned, and the palmar aponeurosis was removed.

Firstly, the presence of the SPA and whether it was a complete arch (complete/incomplete) in case of its presence were evaluated. If the SPA was formed by the UA and superficial palmar branch of the RA, it was evaluated as the "complete" SPA. If it was formed only by the UA, it was evaluated as the "incomplete" SPA. However, according to the definitions in the literature,^[3,4] the SPA formed only by the UA was included in the "complete" SPA category if it reached the first interosseous space. Furthermore, in our study, the number of the common palmar digital arteries emerging from the SPA and the interosseal space of each of these branches were determined. After the measurements of the SPA were completed, the number of branches from the MN and UN, and the interosseal space of each of these branches were recorded. According to the data obtained, the SPA, UN, and MN were classified.

After these procedures, the SPA, UN, and MN were cut from the wrist fold and lifted toward the fingers. The lumbrical muscles and flexor muscle tendons were removed. The DPA was defined, and the number of the metacarpal palmar arteries emerging from the DPA and the interosseal space of these branches were determined. Then, the DPA was classified according to the data obtained.

Results

All of our samples had the SPA. Of the samples, 90% (n = 18) had the "complete" SPA and 10% (n = 2) had the "incomplete" SPA. When the "complete" SPAs were examined, it was revealed that the arch was formed by the UA and superficial palmar branch of the RA (radio-ulnar type) in 83.33% (n = 15) of the samples and only by the UA (ulnar type) in 16.67% (n = 3) of the samples.

When the number of the common palmar digital arteries emerging from the SPA was examined, four branches were observed to emerge from the arch in 80% (n = 16), and five branches emerged from the arch in 20% (n = 4) of the samples. It was detected that 65% (n = 13) of the first branch emerging from the arch ran medially in hand, and 35% (n = 7) ran in the fourth interosseous space. Of the second branch emerging from the arch, 55% (n = 11) was observed to run in the fourth interosseous space and 45% (n = 9) in the third interosseous space. Of the third branch emerging from the arch, 55% (n = 11) was determined to run in the third interosseous space and 45% (n = 9) in the second interosseous space. Of the fourth branch emerging from the arch, 55% (n = 11) was observed to run in the second interosseous space and 45% (n = 9) in the first interosseous space. In the four samples in which the fifth branch emerged, the whole branch was found to run in the first interosseous space.

While classifying the SPA, whether the arch is "complete" or "incomplete," the type of arteries that formed the arch (radio-ulnar type, ulnar type), the number of the common palmar digital arteries given off by the arch, and the course of these branches were taken into consideration, and the arch was gathered under seven main groups [Figures 1-5].

When the innervation of the MN and UN was evaluated, 3.5 + 1.5 fingers innervation was observed in 90% (n = 18), 3 + 1.5 fingers innervation was observed in 5% (n = 1), and 3 + 2 fingers innervation was observed in 5% (n = 1) of the samples.

When 18 samples with 3.5 + 1.5 fingers innervation were examined, the MN was determined to be separated into four branches at the terminal in 94.44% (n = 17) and into five branches in 5.56% (n = 1) of the samples. In the samples in which the MN was separated into four branches, the first branch was observed to course laterally in the hand, the second branch coursed in the first interosseous space, the third branch coursed in the second interosseous space, and the fourth branch coursed in the third interosseous space. In the sample in which the MN was separated into five branches at the terminal, the first branch was observed to run laterally in the hand. The second branch was determined to run in the first interosseous space and innervate the thumb medial. The third branch was determined to run in the first interosseous space, but innervate the lateral of the second finger. The fourth branch emerging from the MN was observed to run in the second interosseous space, and the fifth branch ran in the third interosseous space. In all of the 18 samples with 3.5 + 1.5fingers innervation, the UN was separated into two branches at the terminal. The first branch emerging from the nerve was observed to course in the fourth interosseous space, and the second branch coursed medially in the hand [Figures 6 and 7].

When the sample with 3 + 1.5 fingers innervation was examined, the MN was determined to be separated into three branches at the terminal. The first one of these branches was observed to course in the first interosseous space, the second branch coursed in the second interosseous space, and the third branch coursed in the third interosseous space. In this sample on the right side, it was noted that there were no branches on the lateral side of the thumb. In the same sample, the UN was observed to be separated into two branches at



Figure 1: Schematic classification of the "complete" superficial palmar arch. UA: Ulnar artery, RA: Radial artery, *common palmar digital artery



Figure 3: Types of the "complete" superficial palmar arch that gives off four branches. (a) Type I, (b) Type II, (c) Type III. UA: Ulnar artery, RA: Radial artery, *common palmar digital artery

the terminal. The first branch emerging from the UN was observed to course in the fourth interosseous space, and the second branch coursed medially in hand [Figures 6 and 8].

When the sample with 3 + 2 fingers innervation was examined, the MN was observed to be separated again into four branches. However, when their courses were examined, the branch, which coursed in the third interosseous space, was observed to innervate only the medial of the third finger, i.e., not to innervate the lateral of the 4th finger. The other branches of the MN coursed normally. In this sample, two branches were determined to emerge from the UN. The first one of these branches was observed to run in the fourth interosseous space, and the second branch ran medially in hand. The branch in the fourth interosseous space was determined to be separated into two at the terminal. It was determined that one of these branches innervated the lateral of the fourth finger, the other one continued to run in the fourth interosseous space and innervated the medial of the fourth finger and the lateral of the fifth finger [Figures 6 and 8].

The MN and UN were classified by considering the distribution of the nerves, the number of branches into



Figure 2: Schematic classification of the "incomplete" superficial palmar arch. UA: Ulnar artery, RA: Radial artery, *common palmar digital artery



Figure 4: Types of the "complete" superficial palmar arch that gives off five branches. (a) Type IV, (b) Type V. UA: Ulnar artery, RA: Radial artery, *common palmar digital artery

which these nerves were separated at the terminal, and the course of these branches. Accordingly, the nerves were gathered under four main groups [Figures 6 and 8].

When the DPA samples were examined, four metacarpal palmar arteries were observed to emerge from the arch in all of the samples. It was found that 100% (n = 20) of the first branch ran in the first interosseous space, 95% (n = 9) of the second branch ran in the second interosseous space, and 5% (n = 1) ran in the third interosseous space. Of the third branch emerging from the arch, 95% (n = 19) was observed to run in the fourth interosseous space. Of the fourth branch, 90% (n = 18) was observed to run in the fourth interosseous space, and 10% (n = 2) ran medially in hand.

The DPA was classified according to the number of the metacarpal palmar arteries given off by the arch and the course of these branches. Accordingly, the DPA was gathered under three main groups [Figures 9 and 10].



Figure 5: Types of the "incomplete" superficial palmar arch. (a) Type VI, (b) Type VII. UA: Ulnar artery, RA: Radial artery, *common palmar digital artery



Figure 7: Types of the nerves on the hand with 3.5 + 1.5 finger innervation. (a) Type I, (b) Type II. MN: Median nerve, UN: Ulnar nerve, *terminal branches of the nerves



Figure 9: Schematic classification of the deep palmar arch. UA: Ulnar artery, RA: Radial artery, *metacarpal palmar artery

The frequency and distribution of the SPA types in the right and left hands are presented in Table 1, the frequency and distribution of the MN and UN types in the right and left hands are presented in Table 2, and the frequency and distribution of the DPA types in the right and left hands are presented in Table 3.

Discussion

The vascular structure of the hand, which is a highly complex area, has been investigated in many studies to date. Especially with advances in microsurgery, the vascular structure of the hand has become even more important in reconstructive hand surgery.^[10]

The incidence and variations of the SPA were first defined in the 19th century. In 1897, Jaschtschinski described the "complete" and "incomplete" arch together



Figure 6: Schematic classification of the median nerve and ulnar nerve. MN: Median nerve, UN: Ulnar nerve, *terminal branches of the nerves



Figure 8: (a) Type of the nerves on the hand with 3 + 1.5 finger innervation, Type III. (b) Type of the nerves on the hand with 3 + 2 finger innervation, Type IV. MN: Median nerve, UN: Ulnar nerve, "terminal branches of the nerves



Figure 10: Types of the deep palmar arch. (a) Type I, (b) Type II, (c) Type III. UA: UInar artery, RA: Radial artery, *metacarpal palmar artery

with its different subtypes. Jaschtschinski separated the "complete" SPA into subtypes as the radio-ulnar arch, median-ulnar arch, radio-median-ulnar arch, and ulnar arch.^[23]

The SPA and the frequency of its being complete/ incomplete are presented in Table 4.

Upon examining Table 4, the results obtained from our study were observed to be consistent with the general average. Moreover, according to the table, the results varied in a wide range. For example, the presence of the "complete" SPA ranged between 45.23% and 95%. We think that this variability may be due to differences, such

Table 1: Frequency of the superficial palmar arch types and their distribution in the right and left hands, n (%)					
Туреѕ	Right hand	Left hand	Total		
Туре І	4 (20)	3 (15)	7 (35)		
Type II	4 (20)	1 (5)	5 (25)		
Type III	1 (5)	1 (5)	2 (10)		
Type IV	-	1 (5)	1 (5)		
Type V	-	3 (15)	3 (15)		
Type VI	1 (5)	-	1 (5)		
Type VII	-	1 (5)	1 (5)		

n: Number of extremities

 Table 2: Frequency of the median nerve and ulnar nerve types and their distribution in the right and left hands,

n (%)				
Types	Right hand	Left hand	Total	
Туре І	9 (45)	8 (40)	17 (85)	
Type II	-	1 (5)	1 (5)	
Type III	1 (5)	-	1 (5)	
Type IV	-	1 (5)	1 (5)	

n: Number of extremities

Table 3: Frequency of the deep palmar arch type	es and
their distribution in the right and left hands, <i>n</i>	(%)

Types	Right hand	Left hand	Total	
Туре І	10 (50)	8 (40)	18 (90)	
Type II	-	1 (5)	1 (5)	
Type III	-	1 (5)	1 (5)	
w Number o	C antenna iti an			

n: Number of extremities

as ethnicity, gender, the number of samples, study method, and classification.

The studies conducted on the classification of the arch according to the arteries that form the "complete" SPA and their results are presented in Table 5.

When Table 5 was examined, the "complete" SPA was detected to be usually formed by the UA and superficial palmar branch of the RA (radio-ulnar type). This was followed by the ulnar type (the formation of the SPA only by the UA). Only in the studies conducted by Singh *et al.* and Joshi *et al.*,^[4,24] the ulnar type was found to be more common than the radio-ulnar type. In the studies carried out, the radio-ulnar-median type (the SPA is formed by the UA, the superficial palmar branch of the RA and persistent median artery) was encountered at the lowest rate. When these results were evaluated in terms of types, our study was observed to bear similarities with other studies.

In the studies in Table 5, it was determined that the SPA was classified according to whether it was complete/ incomplete and the type of the arteries forming the arch. However, in our study, in addition to these data, the classification was made by also considering the number of the common palmar digital arteries leaving the arch and the course of these branches [Figures 1 and 2].

The SPA plays an essential role in microsurgery after hand injuries. The collateral circulation of the hand continues even if any artery in the hand is occluded. Plastic surgeons and hand surgeons should know the normal anatomy and variations of these arteries well before performing surgical procedures such as vascular repair and grafting. Since our study was carried out by considering more data on classification, we believe that our study will constitute a basis for future studies and will shed light on surgical procedures.

Although there are various studies on the sensory innervation of the hand in the literature,^[21,29,30] there are no studies involving the classification of nerve distribution

Table 4: The superficial palmar arch and frequency of its being complete/incomplete in the literature					
Researchers, year	Population	Total number of samples (<i>n</i> , number of hands)	Complete superficial palmar arch, <i>n</i> (%)	Incomplete superficial palmar arch, <i>n</i> (%)	
Our study	Turkish	20	18 (90)	2 (10)	
Loukas et al. ^[3]	American	200	180 (90)	20 (10)	
Singh et al. ^[4]	South African	100	92 (92)	8 (8)	
Suma et al. ^[5]	Indian	20	19 (95)	1 (5)	
Gokhroo et al. ^[6]	Indian	200	149 (74.5)	51 (25.5)	
Al-Turk and Metcalf ^[8]	Caucasian	50	42 (84)	8 (16)	
Bilge et al.[10]	Turkish	50	43 (86)	7 (14)	
Tagil et al.[11]	Turkish	20	15 (75)	5 (25)	
Joshi et al. ^[24]	Indian	100	82 (82)	18 (18)	
Sarkar et al.[25]	Indian	42	19 (45.23)	23 (54.76)	
Ottone et al. ^[26]	Argentinian	86	50 (58)	36 (42)	
de Moraes et al. ^[27]	Brazilian	30	18 (60)	12 (40)	

Researchers, year	Population	Radio-ulnar	Ulnar type (%)	Radio-ulnar-median	Ulnar-median type (%)	Ulnar-arcus palmaris
Our study	Turkish	75	15	-	-	-
Loukas <i>et al.</i> ^[3]	American	40	35	6.1	15	3.9
Singh et al. ^[4]	South African	44	46	2	-	-
Al-Turk and Metcalf ^[8]	Caucasian	78	-	2	4	-
Bilge et al.[10]	Turkish	62	14	-	4	6
Joshi et al. ^[24]	Indian	30	66	-	4	-
Sarkar et al.[25]	Indian	89.47	-	-	-	10.52
Ottone et al.[26]	Argentinian	37.1	20.9	-	-	-
de Moraes et al.[27]	Brazilian	36.67	23.34	-	-	-
Madhyastha et al.[28]	Indian	93.75	2.08	-	-	-

Table 5: Classification of the arch according	g to the arteries forming the	"complete"	superficial palmar	arch in the

according to fingers. In our study, the samples were evaluated for innervation by considering the distribution of the MN and UN in hand. Accordingly, 3.5 + 1.5 fingers innervation was observed in 90%, 3 + 1.5 fingers innervation was observed in 5% of the samples, and 3 + 2 fingers innervation was observed in the remaining 5%.

In this study, which we performed by the anatomical dissection method, the MN and UN were classified by considering the distribution of the nerves, the number of branches into which the nerves were separated at the terminal, and the course of these branches [Figure 6]. Our study may be a basic study on this subject. The anatomical dissection method is also known to provide more reliable and accurate results than imaging methods. Therefore, we believe that our study will help to identify and distinguish structures in neurology clinics, neurosurgery, orthopedics, plastic, reconstructive and aesthetic surgery and will provide a basis for studies on the subject.

While there are many studies related to DPA in the literature review,^[3,4,9] this number is very low when the Turkish population is examined.^[12]

In the study conducted by Loukas *et al.*,^[3] the DPA was separated into three types according to the arteries forming the arch. According to this study, if the DPA was formed by the RA and inferior deep palmar branch of the UA, it was defined as "type D-I." If it was formed by the RA and superior deep palmar branch of the UA, it was defined as "type D-II." If it was formed by the RA and both deep palmar branches of the UA, it was defined as "type D-II." If the study, type D-I was found at a rate of 60% (n = 120), type D-II was found at a rate of 30% (n = 60), and type D-III was found at a rate of 10% (n = 20).

In the study performed by Singh *et al.*,^[4] the DPA was separated into five types according to the arteries forming the arch. According to this study, if the DPA was formed by the RA and deep palmar branch of the UA, it was defined as "type G." If it was formed by the RA, deep palmar branch of the UA and interosseous artery, it was defined

as "type H." If the deep palmar branch of the UA was separated into superior and inferior branches and formed an arch with the RA, it was defined as "type I." If the deep palmar branch of the UA was separated into superior and inferior branches, and one of these branches formed an arch with the RA, it was defined as "type J." If the interosseous artery also joined the combination in type "J" and formed an anastomosis with either the arch or the branch of the additional UA, it was defined as "type K." In the study, type G was found at a rate of 72% (n = 72), type H at a rate of 12% (n = 12), type I at a rate of 8% (n = 8), type J at a rate of 4% (n = 4) and type K was found at a rate of 4% (n = 4).

In both studies, the DPA was observed to be classified according to whether the arch was complete/incomplete and the type of the arteries forming the arch.^[3,4] In our study, during classification, as in the SPA, in addition to the data mentioned above, the number of the metacarpal palmar arteries given off by the arch and the course of these branches were also taken into consideration [Figure 9].

There are some limitations of this study since a smaller number of samples. However, in our study, unlike previous studies, more information was given, especially in terms of the course of the branches given by the arches and nerves and typing was made according to the course of these branches.

Conclusions

Understanding the anatomical structure and variations of the arteries and nerves of the hand is particularly important during the surgical reconstruction of the hand in hand injuries. It is known that the lack of knowledge about the anatomy of the palmar region increases the risk of complications during surgical procedures applied to this region. We think that the data obtained on the vascular and neural anatomy of the hand in our study will be included, especially in anatomy education. At the same time, while determining pathologies and variations related to the structures, these data will contribute to the diagnosis and treatment in orthopedics, surgery, radiology and neurology clinics, and other science branches. Thus, it will guide anatomists and other scientists in studies on the subject.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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Multiple Renal Vasculature

Abstract

Variations in the renal vasculature are often encountered in routine cadaveric dissection. In this case, variations were found in the right renal vein (RRV), right renal artery (RRA) and left renal artery (LRA) of a formalin-fixed cadaver. The RRV had two tributaries and the RRA had three branches. The LRA had two branches and was anterior to the left renal vein. Such vascular anomalies are significant clinically.

Keywords: Left renal artery, left renal vein, right renal artery, right renal vein

Introduction

The large renal veins lie anterior to the renal arteries and open into the inferior venacava almost at the right angles. The left renal vein (LRV) three times longer than the right (7.5 cm and 2.5 cm, respectively). For this reason, the left kidney is the preferred side for live donor nephrectomy. The right renal vein (RRV) can be extremely short (<1 cm) such that safe nephrectomy may require excision of a cuff of the inferior venacava. The RRV has no significant collateral drainage like that of the left and cannot be ligated with impunity.

Renal arteries branch laterally from the abdominal aorta just below the origin of the superior mesenteric artery. Both the arteries cross the corresponding crus of the diaphragm at right angles to the aorta. The right renal artery (RRA) is longer and often higher, passing posterior to the inferior venacava, RRV, head of the pancreas and splenic vein.

A single renal artery to each kidney is present in approximately 70% of individuals. The arteries vary in their level of origin and in their caliber, obliquity, and precise relations. In its extra renal course, each renal artery gives off one or more inferior suprarenal arteries, a branch to the ureter, and branches which supply perinephric tissue, the renal capsule and the pelvis. Near the renal hilum each artery



Supernumerary renal arteries are relatively common. They represent persistent fetal renal arteries which grow in sequence from the aorta to supply the kidney as it ascends from the pelvis. Their occurrence is clinically important because a supernumerary artery may cross the pelviureteral junction and obstruct the outflow of urine, producing dilatation of the calyces and pelvis, a condition known as hydronephrosis.^[2]

Aberrant real vessels from the development of kidney it is clear that there may be an aberrant renal artery supplying the kidney. Such artery is usually small and insignificant, but occasionally it may be large and become significant due to the fact that it supplies a segment of the kidney. Such artery is more often seen supplying the lower pole of the kidney. This artery may be associated with hydronephrosis

How to cite this article: Vijaynath V, Ravi V, Mukundan M, Tintu TS. Multiple renal vasculature. J Anat Soc India 2020;69:XX-XX.

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Article Info

Received: 25 December 2019 Accepted: 22 May 2020 Available online: ***

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by compressing the pelvi-ureteric junction or it simply accompanies it, is a point of debate. One must be careful in dividing such aberrant artery as it may cause infarction of the corresponding segment of the renal tissue if it be the only arterial supply to that region.

Aberrant renal artery is more often seen on the left side, and it is more common in females. There may be aberrant renal vein but its significance is much less than aberrant renal artery and it is also less common. Moreover, due to venous collateral circulation, its importance is insignificant.^[3]

Case Report

During routine dissection of a 55-year old male cadaver in the department of anatomy, variations were found in the RRV, RRA, and left renal artery (LRA) simultaneously. There were two tributaries for the RRV, the upper tributary emerged slightly below the middle of the hilum, ran upwards and to the left with an inclination of 45° with the horizontal plane. The upper tributary was about 1.5 cm long and joined the lower tributary at an acute angle. The lower tributary was slender than the upper one and emerged from lower part of the hilum, 1 cm below the exit of the upper tributary. It also ran upwards and to the left, parallel to the upper tributary and joined the upper one at an acute angle after a course of 1.5 cm. The combined wider single RRV, about 1.5 cm wide, continued upward and left for a length of 3 cm and joined the inferior vena cava on its right margin. The tributaries and the single RRV were anterior to the RRA and its branches [Figure 1].

The single RRA emerged from the abdominal aorta under cover of the inferior vena cava and after a course of 1 cm toward the right and in upward direction, divided into three before entering the hilum. The upper branch, 2 cm long, continued upward and to the right, above the upper margin of RRV and entered the hilum at its upper angle. The middle branch ran almost horizontal above the RRV for a distance of 3 cm and entered the hilum just above its center. The lower branch curved downward and entered the hilum near the lower pole. The division of the artery into upper and middle branches was visible just above the middle of the upper border of the RRV [Figure 2].

The LRV was single throughout its length. The vein 1.5 cm wide and 7 cm long, emerged from the middle of the hilum posterior to the entrance of the branches of LRA, ran almost horizontal, dorsal to the branches of LRA. The origin of the superior mesenteric artery lies immediately above the terminal part of the LRV. The terminal part of the LRV curved upward and to the right and entered the inferior vena cava at its left margin, opposite to the entrance of the RRV.

The LRA originated as two branches, upper and lower, from the abdominal aorta, immediately below and slightly to the left of the origin of superior mesenteric artery. The upper branch originated from the left margin and after a very short course of 2 cm divided into two twigs, dorsal and ventral. The origin of branches lay posterior to the LRV. The two branches after a course to the left appeared on the left side of the opening of the lumbar vein to the LRV. The upper branch of LRA then subdivided into ventral and dorsal twigs. The dorsal twig of upper branch ran horizontally to the left and entered the upper angle of the hilum. The ventral twig separated from the common trunk ran slightly downward to the right above the upper margin of the LRV and divided into two. The upper subdivision of the ventral twig of LRA ran horizontally to enter the upper part of the hilum just below the entrance of the upper branch. The lower subdivision of the ventral twig ran almost vertically downward and to the left, crossed the LRV and entered the lower part of the hilum, below the exit of the LRV.



Figure 1: Single right renal vein and its tributaries located in the front of the right renal artery and its branches. (IVC: Inferior vena cava, RRV: Right renal vein with its upper and lower tributaries, PM: Psoas major muscle, U: Ureter)



Figure 2: Divisions of the right renal artery into upper and middle branches visible just above the middle of the upper border of the right renal vein. Inferior vena cava reflected downward. (RRA: Right renal artery with upper, middle and lower branches, RRV: Right renal vein, AA: Abdominal aorta, IVC: Inferior vena cava)

The separate lower branch emerged from the aorta slightly anterior to and below the origin of the common trunk of upper and lower branches and ran upward and anteriorly, wound around the upper margin of LRV and continued downward and to the left crossing anterior to the LRV. The branch opened into the lower pole of the kidney anterior to the renal pelvis [Figure 3].

Discussion

The large renal veins lie anterior to the renal arteries and open into the inferior vena cava almost at right angles. The LRVs are three times longer than the right (7.5 cm and 2.5 cm, respectively). For this reason, the left kidney is the preferred side for live donor nephrectomy. The RRV can be extremely short (<1 cm) such that safe nephrectomy may require excision of a cuff of the inferior vena cava. The RRV has no significant collateral drainage like that of the left and cannot be ligated with impunity.

Renal arteries branch laterally from the abdominal aorta just below the origin of the superior mesenteric artery. Both the arteries cross the corresponding crus of the diaphragm at right angles to the aorta. The RRA is longer and often higher, passing posterior to the inferior vena cava, RRV, head of the pancreas and splenic vein.

A single renal artery to each kidney is present in approximately 70% of individuals. The arteries vary in their level of origin and in their caliber, obliquity, and precise relations. In its extra renal course, each renal artery gives off one or more inferior suprarenal arteries, a branch to the ureter, and branches which supply perinephric tissue, the renal capsule and the pelvis. Near the renal hilum each artery divides into an anterior and a posterior division, and these divide into segmental arteries supplying the renal vascular segments. Accessory renal arteries are common (30% of the individuals) and usually arise from



Figure 3: Left renal artery with its upper and lower branches and their distribution. (ULRA: Upper branch of the left renal artery with its dorsal and ventral twigs, LLRA: Lower branch of the left renal artery, U: Ureter, PM: Psoas major muscle, LRV: Left renal vein)

the aorta above or below (most commonly below) the main renal artery and follow it to the renal hilum. They are regarded as persistent embryonic lateral splanchnic arteries. Accessory vessels to the inferior pole cross anterior to the ureter and may, by obstructing the ureter, cause hydronephrosis. Rarely, accessory renal arteries arise from the coeliac or superior mesenteric arteries near the aortic bifurcation or from the common iliac arteries.^[1]

Supernumerary renal arteries are relatively common. They represent persistent fetal renal arteries which grow in sequence from the aorta to supply the kidney as it ascends from the pelvis. Their occurrence is clinically important because a supernumerary artery may cross the pelvi-ureteral junction and obstruct the outflow of urine, producing dilatation of the calyces and pelvis, a condition known as hydronephrosis.^[2]

Aberrant real vessels from the development of the kidney it is clear that there may be an aberrant renal artery supplying the kidney. Such artery is usually small and insignificant, but occasionally, it may be large and become significant due to the fact that it supplies a segment of the kidney. Such artery is more often seen supplying the lower pole of the kidney. This artery may be associated with hydronephrosis by compressing the pelviureteric junction or it simply accompanies it, is a point of debate. One must be careful in dividing such aberrant artery as it may cause infarction of the corresponding segment of the renal tissue if it be the only arterial supply to that region.

Aberrant renal artery is more often seen on the left side, and it is more common in females. There may be aberrant renal vein, but its significance is much less than aberrant renal artery and it is also less common. Moreover, due to venous collateral circulation, its importance is insignificant.^[3]

The variations in renal vasculature, though very usual in occurrence, are very significant from the surgical point of view. Renal artery variations are of two types: early division and extra renal artery. Branching of the main renal arteries into segmental branches more proximally than the renal hilum level is called early division. Extra renal arteries are of two types: hilum (accessory) and polar (aberrant) arteries. Hilum arteries enter the kidneys from the hilum with the main artery, whereas polar arteries enter the kidney directly from the capsule outside the hilus.^[4]

The LRA in the current case thus showed both the two varieties of the variations. The upper branch showed early division and the lower branch can be considered as an extra renal artery and comes under the polar variety of the extra renal artery. The lower branch can be considered as the aberrant renal artery, and its presence is important from the clinical point of view as this artery may be associated with hydronephrosis by compressing the pelviureteric junction or infarction of the corresponding segment of the renal tissue if it be the only arterial supply to that region.^[3] The RRA showed early division variety of renal artery variation.

In the study conducted by Ozkan *et al.*, including 855 patients, one renal artery was observed feeding both the kidneys in 76% of the patients. More than one renal artery was observed in 24% patients, of which 16% were on the right side and 13% on the left side. In 5% patients, more than one renal artery was present on both sides.^[4] In this case, the extra real artery and early divisions of the renal artery were present in the left side and early division alone in the right side.

Gulas *et al.* has reported that with regard to ethnicity, the incidence of accessory renal arteries fluctuates from 4% in a Malaysian population to 61.5% in a Brazilian population. The frequency is the lowest in the Eastern and Southern Asia (from 4% to 18.4%). In some, not ethnically homogenous populations, wide span of occurrence of accessory renal arteries is described (e.g., American – averaging from 18% to 28.8%). A higher frequency of accessory renal arteries was observed in fetuses compared to adults.^[5]

Dhar and Lal has reported in a study of 40 cadavers that revealed a single main renal artery on either side in 80% of the specimens. Multiple (accessory) renal arteries were observed in 20% of the specimens with unilateral anomaly (15%) being more commonly encountered than bilateral anomaly (5%).^[6]

The right kidney in the present case showed the presence of supernumerary renal vein which is an additional renal vein arising from the renal hilum and draining into the inferior vena cava.

Glodny *et al.*, in a study on hypertension associated with multiple renal arteries on 62 individuals, has concluded that patients with multiple renal arteries constitute a group who have high plasma renin activity and may therefore be prone to develop arterial hypertension.^[7]

Kumar *et al.* has reported a case of an accessory RRV draining into the posterior aspect of the inferior vena cava.^[8] Kumar *et al.* has reported a case of bilateral variations at the hilum involving both renal veins and arteries.^[9] Mukundan *et al.* has reported a case of bilateral double renal vein.^[10]

Satyapal, states that statistically significant changes have been noted in the drainage pattern of right and LRVs. Three hundred and six kidneys were studied and three major types (I, II, and III) of kidney drainage were observed. Type IA consisted of 2 primary tributaries only-an upper and a lower which occurred in 118 (38.6%). It was commonly present on the left side. In addition to upper and lower primary tributaries, a posterior primary tributary was also present in Type IB. This type was observed in 77 (25.2%) cases. In Type IIA, the existence of more than two tributaries, i.e., upper, middle, and lower, were observed. A maximum of five primary tributaries was identified and was noted in 36 (11.8%) cases. In Type IIB, in addition to the primary tributaries present in Type IIA, a posterior primary tributary was also present. Type II B was present in 31 (10.1%) out of the 306 kidneys. Type III consisted of either Types IA, IB, IIA, or IIB along with an additional renal vein or veins. Such variety occurred in 44 (14.4%) cases.^[11]

Conclusion

Variations in the renal vasculature are important from an urologist's perspective. A correct knowledge of this vasculature will help a surgeon while performing renal transplantation and other vascular surgeries. The clinicians dealing with the renal vessels ought to know these variations to keep away from the harm's way.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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Moyamoya Disease in a 29-Year-Old Female of Indian Origin

Abstract

The case presented here is a 29-year-old female patient with complaints of weakness over the left upper and lower limbs for 2 weeks. Digital subtraction angiography of the brain was suggestive of cerebral arteries, showing proliferation of cortical as well as collateral arteries arising from the internal carotid arteries (ICAs). Moyamoya is a progressive steno-occlusive disease at the terminal portions of the bilateral ICAs, with the development of "moyamoya vessels" giving a "Puff of Smoke" appearance on catheter angiography. Cerebral angiography is gold standard for diagnosis and helpful for detecting aneurysm and for surgical planning. There is no standardized surgical strategy for the treatment of this disease. Innumerable revascularization procedures have been employed with the aim of decelerating the progression of disease. Recognition of various symptoms of moyamoya would enable early diagnosis so that it could be treated surgically, as soon as possible, leading to a better neurological outcome.

Keywords: Digital subtraction angiography, moyamoya disease, moyamoya syndrome, neurosurgical interventions, revascularization, stroke

Introduction

Moyamoya disease was first narrated in 1957 by Takeuchi and Shimizu in Japan. Cases have been reported worldwide, but this disease is most common in Japan.^[1] Many cases have been reported in India too.^[2] It is a rare cerebrovascular disease with an incidence of 0.086/100,000 population.^[3] Exact etiology of this disease is not known. It is a progressive steno-occlusive disease. The site of its occurrence is at the terminal portions of the bilateral internal carotid arteries (ICAs). Collateral channels of circulation develop. These small, multiple vessels at the base of the brain are known as "moyamoya vessels." This appearance was observed on catheter angiography and was given the name movamova, which in Japanese means "puff of smoke."^[4,5] Cerebral revascularization surgery leads to favorable outcome of disease progression. Patients are likely to suffer stroke-ischemic or hemorrhagic.^[6] Moyamoya disease generally occurs either in the pediatric age group between 5 and 10 years or in the fourth decade of life.^[7] Generally, ischemic stroke is there in the pediatric group while hemorrhagic stroke in the adults.^[8] Demographically, the ratio

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of females to males is 2:1.^[9] Although moyamoya syndrome is linked with other medical conditions such as arteriosclerosis, Down syndrome, head trauma, autoimmune disease neurofibromatosis type 1, meningitis, or previous radiation therapy, it has the same angiographic appearance as moyamoya disease.^[6]

Case Report

This study was conducted at AIIMS, Jodhpur. Clearance of the institutional ethics committee was taken before commencement of the study. Institutional ethics committee stated "after through consideration accorded its approval on the above project."

A 29-year-old female was admitted to our hospital with complaints of weakness of the left upper and lower limbs for 2 weeks. There was a history of occasional headache and sudden fall thrice in the past 5 months. These sudden falls indicate transient ischemic attacks. There was no history of fever, vomiting, convulsion, and ear discharge. There was no past history of head injury, central nervous system infections, and facial palsy. There was no history of consumption of drugs or noxious substances. She was a full-term vaginal delivery with no complications. There was no history of delayed mile stones. On

How to cite this article: Sharma S, Dixit SG, Khera PS, Garg PK, Nayyar AK, Ghatak S. Moyamoya disease in a 29-year-old female of Indian origin. J Anat Soc India 2020;69:XX-XX.

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Article Info

Received: 23 August 2020 Accepted: 22 May 2020 Available online: ***

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neurological examination, gait was hemiplegic, and tone as well as power was decreased over the left side. Blood, urine, and cerebrospinal fluid investigations were normal. Family history was negative for this disease or any other central nervous system disorder. Hence, after magnetic resonance imaging (MRI) brain, digital subtraction angiography (DSA) of the brain was done. In MRI, many small round and twisted portions of the artery were observed at the base of the brain. DSA was suggestive of significant stenosis of the bilateral ICAs, with the presence of multiple collateral arteries at the base of the brain supplied by vertebral and external carotid arteries, forming "puff of smoke" appearance [Figures 1 and 2].

Discussion

Moyamoya disease is a progressive occlusion of Circle of Willis arteries. This leads to the development of collateral arteries around blocked vessels. This helps in compensating for the blockage. However, the collateral arteries are small and weak. Hence, they are vulnerable to bleeding, thrombosis, and aneurysm. These collateral arteries appear as "puff of smoke" on angiography.^[4] Histopathology of these vessels suggested of thickened intima due to smooth muscle proliferation, as well as tortuous duplicated internal elastic lamina. Abnormalities of the vascular growth factors and inflammatory mediators also have role in the etiology of moyamoya disease.^[10]

Clinical features of moyamoya disease are different in children and adults. In adults, there is history of subarachnoid or intra-parenchymal hemorrhage. However, in children, there is a history of recurrent transient ischemic attacks or infarction of the cerebral artery. Classical symptoms of moyamoya disease are monoparesis, hemiparesis, dysarthria, aphasia, headache, convulsions, and involuntary movements. If diagnosis is done late, then there can be mental impairment too. Two factors on which prognosis of moyamoya disease patients depend are



Figure 1: Digital subtraction angiography of the left common carotid artery lateral (a) and anteroposterior (b) view showing significant attenuation of the supraclinoid internal carotid artery (black arrows) and terminating as ophthalmic artery. Extensive collaterals seen arising from the branches of the external carotid artery (white arrows)

age and type of presentation of disease. Patients with a history of transient ischemic attacks or epilepsy have better prognosis compared to infarctions.^[11]

DSA shows the arteries and collaterals supplying the brain. It is used to visualize the anatomy of the vessels and the presence of aneurysm and hence helps in the confirmation of diagnosis.^[11] In the present case also, DSA showed immense collateral flow through the perforating vessels. Attenuation of the bilateral supraclinoid ICAs with multiple collaterals forming puff of smoke appearance was visualized [Figures 1 and 2].

Antiplatelet agents (including aspirin) and calcium channel blockers are given. Antiplatelet agents are given to prevent clots. Calcium channel blockers provide supportive management of moyamoya disease. Surgery is the best management of moyamoya disease.^[12,13] Moyamoya usually affects the ICA and adjacent anterior and middle cerebral arteries. Therefore, surgeons can use external carotid artery or superficial temporal artery to replace its circulation. The arteries are either sewn directly into brain circulation or placed on the surface of the brain to re-establish new circulation within a few weeks. Revascularization procedures are done at present to increase the perfusion of the hypoxic brain cells. This will ultimately prevent ischemic injury by increasing collateral blood flow.[14,15] Revascularization procedures can be divided into three main groups: indirect (nonanastomotic) bypass techniques, direct (anastomotic) bypass techniques, and combined techniques. Results of direct techniques have been better than indirect and combined techniques.^[16] Angiographic follow-ups have resulted in favorable consequence.

This case emphasizes the significance of considering moyamoya disease as one of the differential diagnoses in patients with a history of recurrent headaches, which cannot be relieved by routine medical treatment.

Conclusion

Early diagnosis and management will lead to favorable results in moyamoya disease patients.



Figure 2: Digital subtraction angiography of the right vertebral artery anteroposterior (a) and lateral (b) view showing extensive collaterals at base of the skull forming puff of smoke appearance (asterisk). Majority of anterior circulation flow is maintained by vertebral artery

DSA helps a lot in diagnosis and henceforth in the management of this disease. This knowledge will be crucial in planning neurosurgical interventions and hence will influence prognosis of moyamoya disease.

Declaration of patient consent

The authors certify that they have obtained all appropriate patient consent forms. In the form the patient(s) has/have given his/her/their consent for his/her/their images and other clinical information to be reported in the journal. The patients understand that their names and initials will not be published and due efforts will be made to conceal their identity, but anonymity cannot be guaranteed.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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