

Volumetric assessment of the sella turcica: a re-evaluation

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Background: The sella turcica volume is widely measured by the Di Chiro-Nelson method. The purpose is to compare the fidelity of a proposed volumetry method vs. the Di Chiro-Nelson method, using computed tomography (CT) images.

Materials and methods: Morphometric examination of 173 CT scans were included, of which 52.6% were female. The mean age was 53.2 ± 17.6 years. Considering the Di Chiro-Nelson method, two measurements were added for each axis in the CT evaluation: length (central, left, and right), width (central, anterior, and posterior), and height (central, left, and right).

Results: The mean measurements were length: central 10.11 ± 1.44 , left 7.45 ± 1.67 , right 7.53 ± 1.59 ; width: central 12.27 ± 2.11 , anterior 10.99 ± 1.92 , posterior 10.10 ± 1.74 ; height: central 7.68 ± 1.38 , left 7.16 ± 1.35 , right 7.40 ± 1.41 . A statistically significant difference between sexes was found only in the anterior width (p = 0.01). Using the proposed method, the volume was 342.2 ± 88.5 and 378.6 ± 113.9 mm³, respectively for females and males (p = 0.02) vs. 476.1 ± 132.4 and 523.8 ± 186.0 mm³ (p = 0.05) using the Di Chiro-Nelson's method.

Conclusions: Women had significantly smaller sella turcica volume than men. This proposed method considers the sella turcica as a not strictly symmetrical structure and indicates reduced variation between the maximum and minimum values, compared to the Di Chiro-Nelson's. Our findings may be useful for reassessment the volume of the sella turcica as the measurements indicate a higher precision. (Folia Morphol 2022; 81, 4: 1014–1021)

Key words: sella turcica, computed tomography, volumetry, Di Chiro-Nelson method

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INTRODUCTION

The sella turcica is a bony structure located in the middle cranial fossa [7]. Its anatomy consists of anterior and posterior clinoid processes and central hypophyseal fossa. The latter is a space formed at the upper surface of the body of the sphenoid bone between the tuberculum sellae and the dorsum sellae [18, 31]. The pituitary gland is a structure that has an important endocrinological function as the central regulator of the endocrine system, located in the deepest point of the sella turcica [17].

There are pathologies and deviations in its development that increase or decrease the size of the pituitary gland. Likewise, these pituitary gland alterations may modify the size and morphometry of the sella turcica, as their development is closely related [27, 35]. The sella turcica abnormalities are caused by individual variations in shape, and asymmetry in normal subjects, pituitary adenomas, hypothyroidism, empty sella syndrome, acromegaly, tumours, and Sheehan syndrome [2, 7, 13, 27]. It also plays a role in craniofacial abnormalities and the decision making and application of surgical interventions [1, 27, 32, 34], as anatomical knowledge is essential during the procedure to avoid damage [35].

Determining the volume of the sella turcica by imaging studies has been used to infer the pituitary gland volume [8, 28] and helps to recognise the normal and anomalous morphology [27]. One of the most widely used volumetry methods is the Di Chiro-Nelson method, in which the length, height, and width are multiplied together and divided in half [8]. However, they utilised radiograph images, and with the improvement of imaging technology, the existing methods have been re-evaluated [21, 27, 35]. We provide a straightforward method, considering the sella turcica as a not strictly three-dimensional symmetrical structure. These measurements play an advantage where magnetic resonance imaging (MRI) is not readily available or contraindicated.

Our objective is to validate a proposed method to determine the volume of the sella turcica and compare it to the Di Chiro-Nelson method using computed tomography (CT) scans. Thus, standardizing the sella turcica volume adds clinical suspicion of pathologies that alter the pituitary gland's size.

MATERIALS AND METHODS

Research design

An observational, retrospective and comparative study was performed. Images were obtained in a case consecutive method from the database of the Radiology and Imaging Department of the University Hospital "José Eleuterio González" of the Universidad Autónoma de Nuevo León. Studies were obtained from adult patients without discrimination for age or gender. Those with a history of fractures, tumours, surgical interventions in the skull base, cancer, craniofacial syndromes, congenital structural abnormalities, or conditions that could affect bone metabolism were excluded. This study adheres to the STROBE guidelines for the report of observational studies [30].

Study technique

All images were acquired using a 64-slice tomograph (General Electric CT99 Light Speed VCT) Software 2978195VCT, with a rotation of 0.4 s helical acquisition, detector coverage of 20 mm, 400 mAs at 120 kV; thickness of cut of 0.625 mm, pitch of 0.53: 1 mm/rot, field of view of 22 to 33 cm. The data obtained were transferred and analysed in a Work Station AW Volume Share2 workstation using multiplanar reformatting with maximum projection intensity and rendering volume. During the measurements, a window range of window width 4000 and window level 1000 was used in a standardised manner in all subjects. The images and measurements were assessed independently by two radiologists with experience in neuroradiology.

The CT scans were measured with Di Chiro-Nelson $[0.5 \times (\text{length} \times \text{width} \times \text{depth})]$ and our proposed method (Table 1). In our proposed method, we added two measurements parallel to each of the three axes (length, height, and width) (Fig. 1). The volume, according to our method, is obtained by the mean of the three measurements of each axis $[0.5 \times [(\text{mean length distances}) \times (\text{mean height distances}) \times (\text{mean width distances})]$. All authors agreed on the bone landmarks, as anatomical variations of the sella turcica shape have been determined.

Ethics approval

This study was previously reviewed and approved by the ethics and research committees of Hospital Universitario "Dr. Jose Eleuterio Gonzalez" of the Universidad Autónoma de Nuevo León under the registration number AH17-00004, certifying that it adheres to the guidelines of the General Health Law on Health Research in Human Beings of our country, as well as international guidelines and the Declaration of Helsinki. Research and ethics

Di Chiro-Nelson method	Description	Current study [2021] method	Description
Length	The distance on the antero- posterior axis of the sella turcica from the back to the tuberculum	Central length	The distance on the anteroposterior axis of the sella turcica from the dorsum to the tubercle
		Left and right length	The distance from each lateral end of the tubercle to the lateral end of the posterior clinoid processes
Height	The line perpendicular to the line that measures the length and that coincides with the deepest portion of the sella turcica	Central height	The line perpendicular to the centre length of the sella to the deepest point of the hypophyseal fossa
		Left and right height	The perpendicular distance of the midpoint of the left and right lengths, respectively to the deepest point of the sella turcica
Width	The lateral-lateral distance of the highest point of the tubercle of the sella turcica	Central width	The lateral-lateral distance of the highest point of the tubercle of the sella turcica
		Anterior width and posterior width	The width of the tubercle and the width of the dorsum of the sella, respectively





Figure 1. Representation of the measurement variables in the proposed method. Measurements represent the variables described in Table 1. Axial slice of computed tomography; A. Central (CL), right (RL), and left length (LL); B. Central (CW), anterior (AW), and posterior width (PW). Sagittal slice of computed tomography; C. Central height (CH); D. Left height (LH); E. Right height (RH).

committees waived the need for further written consent.

Statistical analysis

The sample size calculation was made with a formula for estimating a mean in an infinite population resulting in a sample size of 173 studies. Normality tests were performed using the Kolmogorov-Smirnov test. Central tendency and dispersion data were obtained, expressed as mean and standard deviation for parametric data, and as median and minimum and maximum in nonparametric data. The comparisons between the different groups for the categorical variables were made by Pearson's χ^2 test and for the numerical variables by the two-tailed Student's t test for the parametric data, and Mann-Whitney U test for the non-parametric data. Parametric (Pearson's) and non-parametric (Spearman's) correlations were performed to evaluate the linear relationship between the numerical variables. A p-value of < 0.05 was considered statistically significant. Furthermore, for categorical observations, the inter-observer agreement was assessed by the Cohen's Kappa statistic. A p value of < 0.05 was considered statistically significant. All the statistical analyses were performed in IBM SPSS Statistics for Mac, version 25 (IBM Corp., Armonk, N.Y., USA).

RESULTS

A total of 173 CT scans were included, and 52.6% of the participants were female. The mean age was 53.22 ± 17.62 years (p = 0.73). All sets of inter-observer reliability analyses resulted in substantial reliability (ICC > 0.85 and k > 0.85). The measurements and volume of the sella turcica as means and stratified by sex are shown in Table 2.

Statistically significant differences between sexes were found only in the anterior width (p = 0.01). The comparison of mean volume between sexes was not statistically different using the Di Chiro-Nelson method (p = 0.05) but it was when using the proposed method (p = 0.02) (Table 2). We then compared the mean measurements of both methods and we found a significant difference (p < 0.001).

The differences in standard deviations of the volume were also higher in the Di Chiro-Nelson method (132.41 mm³ in females and 186 mm³ in males) versus the proposed method (88.53 mm³ and 113.89 mm³).

Measure	ment	ent Mean \pm SD Median (minimum–maximum) Females (mean \pm SD) Males (mean \pm S		Males (mean \pm SD)	Р	
Length	Central	10.11 ± 1.44	10.30 (4.3–14.2)	10.11 ± 1.36	10.11 ± 1.54	0.99
	Right	7.53 ± 1.59	7.70 (2.9–11.8)	7.32 ± 1.62	7.68 ± 1.53	0.06
	Left	7.45 ± 1.67	7.40 (2.8–12.2)	7.35 ± 1.77	7.56 ± 1.55	0.42
Width	Central	12.27 ± 2.11	12.30 (7.4–17.5)	12.02 ± 2.12	12.55 ± 2.07	0.09
	Anterior	10.99 ± 1.92	11.00 (5.8–16.2)	10.66 ± 1.87	11.36 ± 1.92	0.01*
	Posterior	10.10 ± 1.74	9.90 (6.0–14.8)	9.95 ± 1.77	10.27 ± 1.70	0.22
Height	Central	7.68 ± 1.38	7.50 (3.4–11.9)	7.50 ± 1.23	7.77 ± 1.51	0.19
	Right	7.40 ± 1.41	7.30 (3.4–11.1)	7.36 ± 1.29	7.44 ± 1.54	0.71
	Left	7.16 ± 1.35	7.10 (3.0–10.6)	7.12 ± 1.34	7.19 ± 1.38	0.72
Volume	Di Chiro's	498.7 ± 161.4	499.6 (90.3–1254.7)	476.1 ± 132.4	523.8 ± 186.0	0.05
	Proposed	359.4 ± 102.7	348.5 (85.6–776.2)	342.2 ± 88.5	378.6 ± 113.9	0.02*

Table 2. Measurements for both sexes and comparison by methods

All measurements in millimetres. Volume in cubic millimetres, *statistical difference between sexes; SD — standard deviation



Figure 2. Box plot demonstrating the values obtained by the Di Chiro-Nelson and the proposed method. The proposed method shows a decrease in the range of the distribution indicating a higher precision of the measurements.

Likewise, the box plot diagram indicates a greater consistency by reducing the variation between the maximum and minimum values, as well as a decrease in the range of the distribution (Fig. 2).

The subjects were stratified according to the age in younger and older than 25 years. A statistically significant correlation is seen between age and the proposed method (r = 0.195) p = 0.01, but not between the Di Chiro-Nelson method (r = 0.130) p = 0.08, in both groups separated by age.

DISCUSSION

Main findings

This study is not the first to use CT to determine the volumetry of the sella turcica. However, we deter-

mined the volume considering it as an asymmetrical structure, in an adult population without pituitary pathology. Our three-dimensional assessment method demonstrates greater consistency and statistically significant differences between sexes. It reports higher fidelity measurements and lower standard deviations than by the Di Chiro-Nelson method, with two-dimensional radiographs (at the publication time, radiographs were one of the standard techniques for pituitary diseases [5]).

Comparison with previous anatomical studies

Other authors have proposed modifications to the Di Chiro-Nelson method (Table 3). Hasan et al. (2016) [9] described the morphology and measurements of the sella turcica in CT images stratified by sex, age, and with global data. No significant differences were found for all linear and area measurements of sella turcica between sexes; however, a gradual increase in the size of sella turcica was observed as age advances [10].

Venieratos et al. (2005) [31] utilised dry skulls and focused on the hypophyseal fossa with a geometrical method as they determined the dimensions of the fossa are smaller than those of the sella, and direct measurement of its volume and depth could lead to errors. However, they measured the total volume of the sella with the Di Chiro-Nelson method, obtaining a range between 460 and 1570 mm³. Hlaing et al. (2012) [11] examined adult crania to observe the bony landmarks and to determine the location of a fossa that may occur in the sellar floor. They described four variations and reported a smooth surface in eight

Author, year, country	Sample	Mean age [years]	Method	Mean volume [mm ³]
Yamada et al., 1976, Japan [33]	570 controls 26 primary hypothyroid patients 34 thyrotoxic patients	Controls: 1–60	Di Chiro-Nelson	$\begin{array}{l} \mbox{Age 1: } 206 \ \pm \ 19 \\ \mbox{Age 1-25: } 530 \ \pm \ 23 \\ \mbox{Age > } 25: \ 554 \ \pm \ 8 \\ \mbox{Hypothyroid: } 1,334.4 \ \pm \ 101.0 \\ \mbox{Thyrotoxic: } 558.3 \ \pm \ 24.7 \end{array}$
Parks et al., 1978, USA [20]	Patients with hypopituitarism: 3	Case 1: 18.6 Case 2: 10.4 Case 3: 6.7	Di Chiro-Nelson	1560 1008 840
Sherif et al., 1989, Libya [26]	Control: 17 Sheehan's: 57	41 ± 8	Statistics volume software and resistor matrix	922 ± 155 565 ± 292
Bakiri et al., 1991, Algeria [4]	Control: 12 Sheehan's: 54	38.3 ± 3.6 40.21 ± 1.22	Height and length on their greatest axis	796 ± 5.6 55.7 ± 2.7
Venieratos et al., 2005, Greece [31]	Dry skulls: 20	NR	Di Chiro-Nelson	835
Pittayapat et al., 2015, Belgium [21]	32 CBCT scans	26.0 ± 21.6	Maxilim [®] software	NR
Hasan et al., 2016, Iraq [10]	71 CT images	33.9	Di Chiro Nelson and three differ- ent heights of the sella turcica	65.3
Yasa et al., 2017, Turkey [35]	177 CBCT scans	11–73	NR	NR
Taner et al., 2019, Turkey [27]	80	F: 26.6 ± 8.6 M: 27.5 ± 9.0	Di Chiro-Nelson	951.3 ± 278.5 1102.0 ± 285.3
Ugurlu et al., 2020, Turkey [29]	Control: 15 Patients with maxillary impacted canines: 73	20.01 ± 6.53	Own method (19 measurements)	NR
Current study, 2021, Mexico	Control: 173 CT scans	$\begin{array}{l} \text{F: 52.8} \pm 18.3 \\ \text{M: 53.6} \pm 17.0 \end{array}$	Proposed method (9 measurements)	F: 342.2 \pm 88.5 M: 378.6 \pm 113.9

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M — male; F — female; NR — not reported; CBCT — cone-beam computed tomography; CT — computed tomography

of the 205 crania. A single posterior fossa was the most common feature, which they described for the first time. We consider these findings to support the reason to measure depth in three different points to have a closer real volume of the sella turcica by taken the anatomical variations into consideration.

Yasa et al. (2017) [35] established the mean distances of length, depth, diameter, and the interclinoid distance of the sella turcica in 177 subjects. No statistically significant difference was found between sexes, but there were statistically significant differences between ages. Our proposed method also identified similar central measurements and a statistically significant correlation between age groups.

Taner et al. (2019) [27] assessed the volume of sella turcica by cone-beam CT in healthy adults. They did not find statistically significant differences between sexes for any Di Chiro-Nelson's method measurements; however, a significant difference was established in volume (p = 0.003), which was

higher in men. Our proposed method showed statistically significant differences between sexes in the anterior width (p = 0.01) and in the volume (p = 0.02).

Pittayapat et al. (2015) [21] developed a high precision system for the sella turcica identification using Maxilim[®] software in 32 subjects. They also focused on orthodontics as the sella point is a reference for the evaluation of the longitudinal growth of patients and their treatment [21]. Our method does not include high precision technology, yet an advantage is that it could be easily replicated with CT images.

Comparison with previous clinical studies

Ugurlu et al. (2019) [29] performed a three-dimensional morphometric analysis of the pituitary fossa with cone-beam CT studies with 19 measurements in subjects with maxillary impacted canines and controls. Only the right sella length differed among the three groups (p < 0.05) [29].

Studies have also determined the volume of the sella turcica in endocrinological diseases. Yamada et al. (1976) [33] compared the size of the sella turcica in normal subjects and patients with primary hypothyroidism and hyperthyroidism. They determined the volume increased with age, and after 25 years of age, remained constant [33].

A small sella turcica size is one predisposing factor that restricts the pituitary blood supply, contributing to Sheehan syndrome [12]. There are studies that compared the volume between patients and controls [4, 26], reporting significantly smaller measurements (p < 0.001) in the patients than in the control subjects [4]. Therefore, it is worthy to establish mean volumes (Table 3), to clinically orient with pituitary disease.

A systematic review by Roomaney and Chetty (2020) [21] determined that the craniofacial morphometry affected by genetic syndromes is likely to be associated with abnormal variations of the sella turcica. Clinicians should be aware of the abnormalities and considering the underlying signs and symptoms for medical referral. They also conclude more high-quality studies are needed, with standardised and objective methods to determine the morphology of the sella turcica.

Di Chiro-Nelson's method was the first used to evaluate pituitary pathology indirectly. Undoubtedly, we can now assess it with MRI and laboratory tests. However, in pathologies where there is a change in the pituitary gland's volume, it is relevant to have a reference for the volume of the sella turcica. Our work could be used to establish a relationship between measurements by CT images and the ones obtained by MRI to standardise an index that supports these diseases' diagnosis.

Limitations of the study

Our method may present inherent imprecision as it has not been validated by other studies or compared to segmentation software methods. Besides, this model attempts to standardize a structure, representing a simplification, given the intrinsic anatomical variability [36]. Further studies are needed to confirm these as there are variations of normal subjects [1, 27], and other abnormal sellar variants, altering sellar volume, such as sellar bridging [9], and variations and morphological types in genetic syndromes (e.g. Williams syndrome) [3]. Also, subjects with dental anomalies and either complete or partial calcification of interclinoid ligament are highly suggestive of a genetic condition [14, 15]. Although there is evidence the sella turcica linear measurements can be used to estimate the pituitary gland size [25], we cannot translate our method. The differences between our morphometric results and those obtained in other studies may be due to the demographic characteristics and imaging methods. We did not perform intraobserver measurements. MRI grants a better understanding of the patient's anatomy and endocrinological diseases [22]. However, it is not always easy to distinguish tumours from haemorrhage and fat packing [16]. CT is less expensive, has a broader distribution, better bone assessment, and it is useful in patients where MRI is contraindicated [24].

Relevance for the clinical practice

The diseases that alter the size of the pituitary are not uncommon [22]. Our results indicate smaller volumes compared to the Di Chiro-Nelson method, and they should be taken cautiously, as a small sella turcica may show upward bulging of the sellar content (one adenomas' indirect sign), leading to potential misinterpretation [19], as well as other normal and abnormal variants that may modify the volume [1, 9, 27].

This study has established a new volume assessment regarding the sella turcica as a variable shape structure enclosing anatomical landmarks. The success in diagnosis and treatment depends on a multicentre management with the integration of internal medicine experts [6]. Nonetheless, the assessment with an anatomical approach provides the clinicians a thorough analysis and better understanding of the related diseases' physiopathology. This is why it is essential to renew the previous methods and consider the sella turcica as a not strictly symmetrical structure. Our findings may be useful to identify the volume of the sella turcica as a component of the integral management of pituitary diseases.

CONCLUSIONS

We determined a volumetric method with greater consistency among measurements and a narrow standard deviation. The Di Chiro-Nelson method has been used for several years. With the new technologies, it was possible to obtain a three-dimensional view of anatomical structures. This is why it is essential to renew the previous methods and consider the sella turcica as a not strictly symmetrical structure. Our findings may be useful to identify the volume of the sella turcica as a component of the integral management of pituitary diseases.

Conflict of interest: None declared

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