Expanded endoscopic endonasal transsphenoidal approach to determine morphological characteristics and clinical considerations of the cavernous sinus venous spaces

Guowen Zhan1, Shanshan Guo1, Honglei Hu1, Jianchun Liao2, Ruishan Dang3 & Youxiong Yang1

The study aimed at investigating the morphological characteristics and interconnected regularities of the cavernous sinus (CS) venous spaces using an expanded endoscopic endonasal transsphenoidal approach. Surgical dissections were performed for 15-colored silicon-injected human head specimens. The CS venous spaces were examined for their morphological and clinical characteristics using an expanded endoscopic endonasal transsphenoidal approach. The intracavernous course of the internal carotid artery (ICA) divided the CS venous spaces into four interconnected virtual compartments: medial, anteroinferior, posterosuperior, and lateral. The CS venous spaces had peculiar morphological characteristics; the medial compartment was C-shaped while the anteroinferior compartment resembled a boat’s bow. The mean distances from the medial border of the inferior horizontal segment of cavernous ICA to the mid-line of the pituitary gland (PG) were 6.07 ± 1.61 mm (left) and 5.97 ± 1.89 mm (right); the mean distances from the medial border of the subarachnoid segment of cavernous ICA to the mid-line of the PG were 5.77 ± 1.16 mm (left) and 5.63 ± 1.17 mm (right); the mean distances from the medial border of the anterior vertical segment of cavernous ICA to the mid-line of the PG were 10.27 ± 1.74 mm (left) and 10.47 ± 1.90 mm (right). Morphological characteristics and the knowledge of the interconnected regularities of the CS venous spaces may help surgeons accurately locate the neurovascular structure, and thus may contribute to the effective prediction of tumor invasion and extension during endoscopic CS surgery.

Expanded endoscopic endonasal approaches have become increasingly popular in the last decade as they facilitate access to virtually any region of the ventral skull base including the anterior cranial fossa, parasellar region, clival region, and craniovertebral junction. Advances in endoscopic endonasal techniques, image-guided systems, neuronavigation, and customized surgical instruments have enabled surgeons to treat skull base lesions with lesser morbidities. The surgeons benefit from the advantages of this approach which include direct access to the ventral skull base by avoiding brain and brainstem retraction, near-field magnification, better surgical field illumination, and minimal manipulation of neurovascular structures.

Advances in the expanded endoscopic endonasal transsphenoidal approach have provided for a new route to access the lesions in anatomically complicated structures in the skull base, which are close to the internal carotid arteries, pituitary gland, abducens nerve, and CS. However, because of the depth of the target location and the complexity of the critical regional anatomy, the exposure and resection of these tumors pose a technical challenge.

1Department of Otolaryngology-Head and Neck Surgery, Ningbo Yinzhou Second Hospital, 1st North Qianhe Road, Yinzhou District, Ningbo 315100, Zhejiang, People’s Republic of China. 2Department of Otolaryngology-Head and Neck Surgery, the Affiliated Changzheng Hospital of Second Military Medical University, Shanghai 200003, People’s Republic of China. 3Department of Human Anatomy, Second Military Medical University, Shanghai 200433, People’s Republic of China. *email: yyy929929@163.com
challenge to the neurosurgeons and otolaryngologists. Space-occupying lesions within this region may distort normal anatomy, and thus stereotactic image guidance can be misleading. Hence, there is a need for more reliable anatomical landmarks for surgery. While the different walls of the CS, the intracavernous ICA, and its branches are well described, morphological and clinical characteristics of the CS venous spaces, remain relatively unknown.

Our previous study has investigated the relationship between the morphological characteristics of the sphenoid sinus and endoscopic localization of the cavernous sinus, and we believe that the opticocarotid recess (OCR) and C-shaped ICA in the lateral wall of the sphenoid sinus are the 2 reliable anatomic landmarks in the intraoperative location of the parsellar region of CS. While in this study, using an expanded endoscopic endonasal transsphenoidal approach, surgical dissections were performed on adult cadaver heads to examine the morphological and clinical characteristics of the four compartments of the CS venous spaces. These may have implications for safe surgical procedures with minimal invasiveness.

Materials and methods
This study was approved by the Ethics Committee of Ningbo Yinzhou Second Hospital (Ningbo, China, 2022003) per the Declaration of Helsinki of 1964 (revised, Hong Kong, 1989). The human cadavers used in the study were supported by the department of Human Anatomy, Second Military Medical University, and informed consents were obtained from all donors or next kin in accordance with relevant guidelines and regulations. Specimens were obtained from 15 randomly selected adult formalin-processed cadaver heads. Red latex was injected into the internal carotid arteries and blue latex in the internal jugular veins. All anatomical dissections were performed using an expanded endoscopic endonasal transsphenoidal approach in the operating room of the Second Yinzhou Hospital. A rigid endoscope (Karl Storz GmbH and Co., Tuttingen, Germany), 4 mm in diameter, 18 cm in length, equipped with 0° lenses, was used. A fiberoptic cable connected the endoscope to a light source, and a camera fitted with high-definition second-generation sensors. The video camera was connected to a monitor which displayed a high-definition image. Cadaveric specimens were placed in a supine position; the head was placed at 10 to 15 degrees adducted toward the left shoulder; in a neutral position. Using the bilateral nostril technique, we introduced the endoscope and instruments through the two nostrils. The dissection was performed in four stages as follows: exposure of the anterior, posterior, and lateral walls of the sphenoid sinus; exposure of the medial and anteroinferior walls of the CS; exposure of the medial and anteroinferior compartments of the CS venous spaces; exposure of the posterosecond posterior and lateral compartments of the CS venous spaces. A custom-made 1 mm × 1 mm grid was designed on a computer and printed on a transparent acetate sheet. This was used to measure the distances from bilateral cavernous ICAs to the mid-line of the PG. Statistical analyses were carried out using the GraphPad Prism 5 (Version 5.01, GraphPad Software, Inc., USA), and the data were presented as mean ± standard deviation (SD). Analysis for statistical significance was performed using the paired t-test and differences were considered as statistically significant for p < 0.05.

Results
Stage 1: Exposure of the anterior, posterior, and lateral walls of sphenoidal sinus. The middle and superior turbinates were laterally retracted, and the sphenoid Ostia were bilaterally identified. Septal mucosal cuts were made 20 mm in front of the ostium, 1 mm above the palate, and 2–3 mm below the cribriform plate (until they met anteriorly), for nasoseptal flap harvesting (Fig. 1).

The sphenoidal sinus at the center of the cranial base was surrounded by numerous neurovascular structures. The bony protuberances of the posterior and lateral walls of the sphenoidal sinus were exposed using complete posterior ethmoidectomy, and wide anterior sphenoidotomy. (Fig. 2).

Stage 2: Exposure of the medial and anteroinferior walls of the CS. When the bone overlying the sphenoidal sinus and sellar floor was removed, the dural layer was exposed and it enveloped the anterior surface of the pituitary gland and the medial wall of the CS. The layer in front of the pituitary gland consists of loosely attached two-layered dura; the external layer called the meningeal dural layer is thicker and extends outwards to the cribriform plate (unti they met anteriorly). This was used to measure the distances from bilateral cavernous ICAs to the mid-line of the PG. Statistical analyses were carried out using the GraphPad Prism 5 (Version 5.01, GraphPad Software, Inc., USA), and the data were presented as mean ± standard deviation (SD). Analysis for statistical significance was performed using the paired t-test and differences were considered as statistically significant for p < 0.05.

Stage 3: Exposure of medial and anteroinferior compartments of the CS venous spaces. Based on their relationship with the cavernous ICA, four CS compartments were classified as follows: medial, anteroinferior, posterosecond, and lateral compartments. The layers covering the medial and anteroinferior walls of the CS were resected to expose the medial and anteroinferior compartments of the CS venous spaces. The medial compartment of the CS venous space was located between cavernous ICA and pituitary gland, and was superiorly limited by the distal tail, and inferiorly by the inferior horizontal psICA. This compartment lacked nerves, and thus represented a medial surgical corridor owing to a minimal risk of neural damage. Anterio–posteriorly, the medial compartment was C-shaped (70%) and non-C-shaped (30%). The medial C-shaped compartment,
anteroinferiorly communicates with the anteroinferior compartment, laterally with the lateral compartment, and posteriorly with the posterosuperior compartment; it also medially crosses the anterior and inferior intercavernous sinuses to the opposite compartment (Fig. 4). The superior hypophyseal artery arising from the ICA located above the medial compartment ran in the preinfundibular space and provided the maximum blood supply to the infundibulum, optic chiasm, and proximal optic nerve (Fig. 5c).

A custom-made 1 mm × 1 mm grid was used to measure the distances from the bilateral cavernous ICAs to the mid-line of the PG and the data are presented as mean ± SD. The mean distance A, from the medial border of the lower cavernous ICA (inferior horizontal segment) to the mid-line of the PG was 6.02 ± 1.73 mm, the mean distance C; from the medial border of the upper cavernous ICA (subarachnoid segment) to the mid-line
Figure 3. Endoscopic endonasal view of the ICAs, CSs and ICSs. (a) After the bone overlying the sphenoidal sinus and sellar floor is removed, the dural layer is shown. (b) The layer and latex covering the anterior portion of the pituitary gland and the medial wall of the CS, are resected to expose the CS compartments. ICSs are venous interconnections between the bilateral CSs. C clivus, ON optic nerve, psICA parasellar segment of the ICA, pcICA paracalval segment of the ICA, PG pituitary gland, DR distal ring, PR proximal ring, aICS anterior intercavernous sinus, ilCS inferior intercavernous sinus, Med.Com. medial compartment of the CS, Ant.Inf.Com. anteroinferior compartment of the CS. *Opticocarotid recess.

Figure 4. Measurements of the distances between the bilateral cavernous ICAs to the mid-line of the PG in endoscopic view and the medial compartment resembles C-shape (the black dotted line), A (the red dotted line) represents the distance from the medial border of the upper cavernous ICA (subarachnoid segment) to the mid-line of the PG, B (the yellow dotted line) is the distance between the medial border of the anterior vertical segment of cavernous ICA to the mid-line of the PG, C (the blue dotted line) marks the distance from the medial border of the lower cavernous ICA (inferior horizontal segment) to the mid-line of the PG, the medial C-shaped compartment communicates with the anteroinferior compartment anteroinferiorly, the lateral compartment laterally and the posterosuperior compartment posteriorly, also crosses the anterior and inferior intercavernous sinus medially to the opposite compartment (arrows). PG pituitary gland, C clivus, ON optic nerve, lhsICA inferior horizontal segment of the ICA, ssICA subarachnoid segment of the ICA, pcICA paracalval segment of the ICA, *Opticocarotid recess.
of the PG was $5.70 \pm 1.15$ mm; the mean distance $B$ from the medial border of the anterior vertical segment of cavernous ICA to the mid-line of the PG was $10.37 \pm 1.79$ mm (Fig. 4). There were no significant differences between the left and the right sides. All measurements are displayed in Table 1.

The relationship between the lateral surface of the pituitary gland (PG) and the inferior horizontal segments of the internal carotid artery (ihsICA) was stratified into three types: contiguity (40%, 12/30), separation (26.7%, 8/30), and indentation (33.3%, 10/30). Notably, the relationship between the lateral surface of the PG and bilateral ihsICAs were not always consistent, (20%, 3/15), as shown in Table 2 and Fig. 5.

The medial wall of the CS was removed. The anteroinferior compartment which was superior to the bottom of CS and inferior to the proximal dural ring of the CS venous space resembled a boat’s bow. It was anteriorly limited.

Figure 5. The relationship between the lateral surface of the pituitary gland (PG) and the inferior horizontal segments of the internal carotid artery (ihsICA). (a) the bilateral ihsICAs (the red dotted line) are attached contacts with the lateral surface of the PG (the blue dotted line) corresponding to contiguity. (b) the bilateral ihsICAs are separated from the lateral surface of the PG corresponding of separation. (c) the bilateral ihsICAs are extended to the lateral surface of the PG corresponding to indentation. (d) the right ihsICA is separated from the lateral surface of the PG whereas the left one is close contact. ON optic nerve, OC optic chiasm, OA ophthalmic artery, SHA superior hypophyseal artery, PG pituitary gland, C clivus, pcICA paraclival segment of the ICA, his inferior horizontal segment of the ICA, avs anterior vertical segment of the ICA, cs clinoidal segment of the ICA, ss subarachnoid segment of the ICA.
by the CS anteroinferior wall, and posteriorly by the anterior surface of the anterior vertical psICA (Fig. 6b). The first nerve we encountered in the anteroinferior compartment around the ICA was the sympathetic plexus; it extended from the pcICA to the ihsICA. The sympathetic nerve was located medial to the abducens nerve with an oblique trajectory. It ran from the surface of the ICA and joined the ophthalmic nerve, with a more horizontal trajectory in this segment (Fig. 6c,d). We identified that the nasal mucosa was bilaterally elevated from the vomer along the inferior wall of the sphenoid sinus up to the vidian canal. The vidian nerve is formed by the union of the greater superficial petrosal nerve and the deep petrosal nerve and is a useful landmark to access the lacerum segment of the ICA (Figs. 2, 6a).

Table 1. Measurements of the distances between bilateral cavernous ICAs to the mid-line of the PG in endoscopic view. Distance A represents the distance from the medial border of the subarachnoid segment of the ICA to the mid-line of the PG. Distance B is the distance between the medial border of the anterior vertical segment of cavernous ICA to the mid-line of the PG. Distance C marks the distance from the medial border of the inferior horizontal segment of the ICA to the mid-line of the PG.

| Specimen no. | Distance A (mm) | Distance B (mm) | Distance C (mm) |
|--------------|----------------|----------------|----------------|
|              | Left Right     | Left Right     | Left Right     |
| 1            | 4 4.5          | 9 8            | 4.5 4          |
| 2            | 5.5 6.0        | 13 13          | 6.5 6.5        |
| 3            | 5.5 5.0        | 11.5 11.5      | 5 4           |
| 4            | 7 5            | 10 10.5        | 6.0 6.5        |
| 5            | 6.5 7          | 7 7            | 5 4           |
| 6            | 3 3            | 9 8            | 3.5 4.0        |
| 7            | 6 6.5          | 12 13          | 5.5 5          |
| 8            | 5 6            | 11.5 11.5      | 6.5 6.5        |
| 9            | 7.0 6.0        | 12 13          | 7.5 8.5        |
| 10           | 6.5 5.5        | 9 10           | 5.0 4.5        |
| 11           | 7 8            | 12 12          | 10.5 10.5      |
| 12           | 6.5 6.5        | 8 10           | 5.5 5          |
| 13           | 5 5            | 11 9           | 6.5 7.5        |
| 14           | 6.5 5.5        | 10 10.5        | 6.5 6.5        |
| 15           | 5.5 5          | 9 10           | 7 6.5          |
| Mean ± SD    | 5.77 ± 1.16    | 5.63 ± 1.17    | 10.27 ± 1.74   |
| Total        | 5.70 ± 1.15    | 10.37 ± 1.79   | 6.02 ± 1.73    |

Table 2. The relationship between the lateral surface of the pituitary gland (PG) and the inferior horizontal segments of the internal carotid artery (ihSICA).

| Specimen no. | The relationship between the lateral surface of PG and ihSICA |
|--------------|-----------------------------------------------------------|
|              | Left Right                                                |
| 1            | Contiguity Contiguity                                     |
| 2            | Indentation Indentation                                   |
| 3            | Indentation Indentation                                   |
| 4            | Separation Separation                                     |
| 5            | Separation Indentation                                    |
| 6            | Indentation Indentation                                   |
| 7            | Indentation Indentation                                   |
| 8            | Contiguity Contiguity                                     |
| 9            | Contiguity Separation                                     |
| 10           | Contiguity Contiguity                                     |
| 11           | Separation Separation                                     |
| 12           | Separation Separation                                     |
| 13           | Indentation Contiguity                                    |
| 14           | Contiguity Contiguity                                     |
| 15           | Contiguity Contiguity                                     |
Stage 4: Exposure of posterosuperior and lateral compartments of the CS venous spaces. The oculomotor, trochlear, ophthalmic (first trigeminal division), and maxillary (second trigeminal division) nerves are embedded in the reticularis membrane forming the lateral compartment of the CS; the abducens nerve, the intracavernous carotid artery, and the surrounding sympathetic plexus have a true intracavernous course. The oculomotor nerve, trochlear nerve, the first division of the trigeminal nerve, and abducens nerve formed a nerve bundle at the superior orbital fissure. Gentle medial mobilization of the ICA genu is required to gain access to the lateral compartment of the CS venous. The lateral compartment is lateral to the anterior vertical psICA and medial to the CS lateral wall. It includes ILT and the 4 triangles (the supratrochlear, infratrochlear, anteromedial, and anterolateral triangles).
The ILT, also known as the inferior artery of the cavernous sinus, arises laterally from the inferior horizontal segments of the ICA, III oculomotor nerve, IV trochlear nerve, VI abducens nerve, ILT inferolateral trunk, Sup.branch superior branch of the ILT, Ant.branch anterior branch of the ILT, Pos.branch posterior branch of the ILT.

Figure 7. Endoscopic view of the ILT and its branches, the ILT originates from the lateral aspect of the inferior horizontal segments of the ICA, III oculomotor nerve, IV trochlear nerve, VI abducens nerve, ILT inferolateral trunk, Sup.branch superior branch of the ILT, Ant.branch anterior branch of the ILT, Pos.branch posterior branch of the ILT.

Discussion
The CS is a valveless, dural-lined, venous plexus located near the center of the head, on either side of the sella, pituitary gland, and sphenoid sinus. The cavernous segment of the ICA lies medial to the CS and is surrounded by a periarterial sympathetic plexus. Courses of Cranial nerves III, IV, V1, and V2 run from top to bottom within the lateral dural walls of each CS. Cranial nerve VI lies immediately inferolateral to the ICA and is the only cranial nerve located within the venous sinusoids of the CS. The CS extends frontally from the superior orbital fissure and extends at the back of the area lateral to the dorsum sellae.

The classification of the CS venous spaces by Harris and Rhoton in 1976, is based on differentiated posterosuperior, anteroinferior, and medial venous spaces with the ICA. Juan C Fernandez-Miranda et al. classify the four CS compartments based on their spatial orientation to the cavernous ICA: superior, posterior, inferior, and lateral compartments. Cavallo LM et al. report that the meningeal wall of the CS has three weak points for tumor invasion and extension: the venous plexus around the SOF, the loose texture of the medial wall around the...
The medial wall of the cavernous sinus is formed by a single, continuous, and thin layer of the dura, while the lateral and the superior walls are two-layered (meningeal and endosteal). The thin dura in the medial wall may be incomplete or absent. Most nonfunctioning pituitary adenomas extending into the cavernous sinus, are neither aggressive nor invasive, the lesions grow through the weak points of the walls of the cavernous sinus. While Ceylan et al. showed the integrity of the medial wall and its ultrastructural integrity with reverse transcription polymerase chain reaction (RT-PCR). Type IV collagen was considered to be an important factor in the progression of adenoma and invasion. The medial compartment has vital clinical utility because pituitary adenomas often invade the medial compartment of the CS through the venous connections by anteroinferior, posterior, and lateral extensions. Therefore, we classified the CS into four walls: medial, anteroinferior, posterosuperior, and lateral walls; and correspondingly the CS venous spaces into four compartments: medial, anteroinferior, posterosuperior, and lateral compartments.

Evidence of venous bleeding from the cavernous sinus is different. Carotid artery injury can occur when bone removal over the CS and along Meckel cave is being performed. Careful use of Kerrison rongeurs and drills is mandatory. Torquing or twisting motion with rongeurs should be avoided because such maneuvers can lead to shards of bone being driven into the vessel wall resulting in laceration or avulsion injuries. Care should be exercised when working on the medial compartment of the CS venous space as follows:

1. During the dural opening of the upper and lower parts of the C-shaped medial compartment, the prominent anterior or inferior intercavernous sinuses pose limitations and may contribute to incomplete tumor resections (Figs. 3b and 6b). The bipolar forceps are used to reduce the risk of preoperative bleeding after an initial small dural incision is made.
2. The superior hypophyseal artery is more likely to arise from the clinoidal segment of the ICA proximal to the DDR (Fig. 5c). It is remarkably difficult to avoid its accidental damage at the origin, especially with bipolar coagulation, while operating on the upper end of the C-shaped medial compartment.
3. It is critical to identify the relationship between the lateral surface of the PG and the ihsICA before the surgery, to design the surgical procedure accordingly (Fig. 5). The indentation group may alter the course of the ICA. Vascular injury is one of the major concerns, especially due to the diverse relationship between the lateral surface of the PG and bilateral ihsICAs; this was found in 20% of the total specimens.

We speculated that distance measurements from the bilateral cavernous ICAs to the mid-line of the PG may benefit and support surgeons in extended approaches, and thus improve resection rates of lesions in this region. Thus, the distance from the bilateral cavernous ICAs to the mid-line of the PG was determined; these results are shown in Table 1. The difference between distances A (the distance from the medial border of the ssICA to the mid-line of the PG) and C (the distance from the medial border of the ihsICA to the mid-line of the PG) was not significant (5.70 ± 1.15 mm vs. 6.02 ± 1.73 mm); the distance from the medial border of the anterior vertical segment of cavernous ICA to the mid-line of the PG was 10.37 ± 1.79 mm. If possible, vascularized reconstruction could be used (often the nasoseptal flap) to cover and protect the ICA when there is vascular exposure.

The anteroinferior compartment of the CS venous space resembled a boat’s bow, and its surgery requires the removal of the bone that covers the anteroinferior wall of the CS. Sympathetic fiber bundles travel along the surface of the carotid as it emerges from the foramen lacerum. Some of the bundles join the abducens nerve within the sinus and are ultimately distributed to the ophthalmic branch of the trigeminal nerve, which sends sympathetic
fibers to the pupillodilator through the long ciliary nerves and the ciliary ganglion (Fig. 6c,d). The vidian nerve which consistently marks the level of the lacerum segment of the ICA in the petrous bone is one of the most important surgical landmarks to navigate the anteroinferior compartment[2].

Our study showed that expanded endoscopic endonasal transphenoidal approach provided excellent views of the medial and anteroinferior compartments of the CS venous spaces, with peculiar morphological characteristics. The medial compartment was C-shaped while the anteroinferior compartment of the CS resembled a boat's bow. These may help surgeons to accurately locate the neurovascular structures during endoscopic CS surgery. However, for the posterosuperior compartment of the CS venous spaces, which is frequently obliterated by the ICA, the endoscopic supraorbital extradural approach may offer better exposures[33]; the lateral compartment of the CS venous spaces is very narrow, thus, the transethmoid-pterigoid-sphenoidal endoscopic approach[5] or endoscopic transorbital approach[44] may be better for maneuverability. Several studies indicate that the posterior compartment may be an important venous compartment as well[26,35]. Especially recently, isolated posterior compartment involvement has also been reported in clinical series[46]. However, the posterior compartment has not been studied in this article.

The expanded endoscopic endonasal transphenoidal approach should be performed only by experienced and well-trained neurosurgeons and otolaryngologists. This can be done by mastering the use of the endoscope in several endoscopic endonasal surgeries and endoscopic cadaver dissections. We believe that morphological characteristics like the one we found and those from other studies have potential clinical applicability in improving the safety of endoscopic skull base surgeries.

Data availability

The authors confirm that all data generated or analysed during this study are included in this published article and its supplementary materials.

Received: 22 January 2022; Accepted: 26 September 2022
Published online: 06 October 2022

References

1. Abuzayed, B. et al. Expanded endoscopic endonasal approach to the clival region. J. Craniofac. Surg. 21, 245–251 (2010).
2. Taniguchi, M. & Kohmura, E. Endoscopic endonasal removal of laterally Expanded clival chordoma using side-viewing scopes. Acta Neurochir. (Wien). 154, 627–632 (2012).
3. Cavallo, L. M. et al. The expanded endoscopic endonasal approach to the clivus and cranio-vertebral junction: Anatomical study. Childs Nerv. Syst. 23, 665–671 (2007).
4. Kamat, A. et al. Reconstructive challenges in the Expanded endoscopic transclival approach. J. Laryngol. Otol. 129, 468–472 (2015).
5. Shin, M. et al. Endoscopic transphenoidal anterior petrosal approach for locally aggressive tumors involving the internal auditory canal, jugular fossa, and cavernous sinus. J. Neurosurg. 126(1), 212–221 (2017).
6. de Notaris, M. et al. Endoscopic endonasal transclival approach and retrosigmoid approach to the clivus and petroclival regions. Neurosurgery 65(suppl 6), 42–50 (2009).
7. de Lara, D. et al. Endonasal endoscopic approaches to the paramedian skull base. World Neurosurg. 82(suppl 6), 121–129 (2014).
8. Pamias-Portalatin, E. et al. Endoscope-assisted contralateral transtemporal approach to the clivus and the hypoglossal canal: technical case report. J. Neurosurg. 22, 1–7 (2018).
9. Inoue, A. et al. Role of denosumab in endoscopic endonasal treatment for juvenile clival giant cell tumor: A case report and review of the literature. World Neurosurg. 91(674), e1–e6 (2016).
10. Saito, K., Toda, M., Tomita, T., Ogawa, K. & Yoshida, K. Surgical results of an endoscopic endonasal approach for clival chordomas. Acta Neurochir. (Wien). 154, 879–886 (2012).
11. Hong Jiang, W. et al. Endoscopic resection of chordomas in different clival regions. Acta Otolaryngol. 129, 71–83 (2009).
12. Fernandes Cabral, D. T. et al. Endoscopic endonasal transclival approach for resection of a pontine glioma: surgical planning, surgical anatomy, and technique. Oper. Neurosurg. (Hagerstown) 12, 589–599 (2018).
13. Ciporen, J., Lucke-Wold, B., Dogan, A., Cetas, J. S. & Cameron, W. E. Dual endoscopic endonasal transphenoidal and preuncal transorbital approaches for clipping of the cavernous carotid artery: A cadaveric simulation. J. Neurol. Surg. B Skull Base 77, 485–490 (2016).
14. Mendelson, Z. S., Patel, A. A., Eloy, J. A. & Liu, J. K. Endoscopic palliative decompression of the cavernous sinus in a rare case of a metastatic renal cell carcinoma to the clivus. Br. J. Neurosurg. 29, 430–431 (2015).
15. Nakamura, S., Kawamata, T., Kobayashi, T. & Horii, T. Clival inflammation with cavernous sinus thrombophlebitis and orbital subperiosteal abscess–case report. Neurol. Med. Chir. (Tokyo) 50, 168–171 (2010).
16. Yildirim, A. E., Divanaloglu, D., Cetinalp, N. E., Ozhamam, E. & Belen, A. D. Endoscopic endonasal transphenoidal and preuncal transorbital approaches for clipping of the cavernous carotid artery: A cadaveric simulation. J. Neurol. Surg. B Skull Base 77, 485–490 (2016).
17. Alfieri, A. & Jho, H. D. Endoscopic endonasal transsphenoidal approach to the clivus: An anatomic study. Neurosurgery 48, 827–836 (2001).
18. Yasuda, A., Campero, A., Martins, C., Rhoton, A. L. & Ribas, G. C. The medial wall of the cavernous sinus: Microsurgical anatomy. Neurosurgery 55(1), 179–189 (2004) (discussion 189–190).
19. Munawar, K. et al. Cavernous sinus lesions. Clin. Imaging 68, 71–89 (2020).
20. Campero, A., Campero, A. A., Martins, C., Yasuda, A. & Rhoton, A. L. Surgical anatomy of the dural walls of the cavernous sinus. J. Craniofac. Surg. 17(6), 746–750 (2010).
21. Yasuda, A. et al. Microsurgical anatomy and approaches to the cavernous sinus. Neurosurgery 56(1 Suppl), 4–27 (2005) (discussion 4–27).
22. Chung, B. S., Chung, M. S. & Park, J. S. Six walls of the cavernous sinus identified by sectioned images and three-dimensional models: Anatomic report. World Neurosurg. 84(2), 337–344 (2015).
23. Harris, F. S. & Rhoton, A. L. Anatomy of the cavernous sinus. A microsurgical study. J. Neurosurg. 45, 169–180 (1976).
24. Fernandez-Miranda, J. C. et al. Cavernous sinus compartments from the endoscopic endonasal approach: Anatomical considerations and surgical relevance to adenoma surgery. J. Neurosurg. 129(2), 430–441 (2018).
27. Cavallo, L. M. et al. Endoscopic transnasal approach to the cavernous sinus versus transcranial route: Anatomic study. *Neurosurgery* **56**(suppl 2), 379–389 (2005).
28. Ceylan, S. et al. Microsurgical anatomy of membranous layers of the pituitary gland and the expression of extracellular matrix collagenous proteins. *Acta Neurochir.* (Wien) **153**(12), 2435–2443 (2011) (discussion 2443).
29. Lobo, B., Zhang, X., Barkhoudarian, G., Griffiths, C. F. & Kelly, D. F. Endonasal endoscopic management of parasellar and cavernous sinus meningiomas. *Neurosurg. Clin. N. Am.* **26**, 389–401 (2015).
30. Chenin, L., Toussaint, P., Lefranc, M., Havet, E. & Peltier, J. Microsurgical anatomy of the inferior intercavernous sinus. *Surg. Radiol. Anat.* **43**(2), 211–218 (2021).
31. Truong, H. Q. et al. Surgical anatomy of the superior hypophyseal artery and its relevance for endoscopic endonasal surgery. *J. Neurosurg.* **131**(1), 154–162 (2018).
32. Rhoton, A. L. The cavernous sinus, the cavernous venous plexus, and the carotid collar. *Neurosurgery* **51**(4 Suppl), S375–410 (2002).
33. Komatsu, F., Komatsu, M., Inoue, T. & Tschabitscher, M. Endoscopic supraorbital extradural approach to the cavernous sinus: A cadaver study. *J. Neurosurg.* **114**, 1331–1337 (2011).
34. Caklili, M., Emengen, A., Cabuk, B., Anik, I. & Ceylan, S. Endoscopic transorbital approach to the cavernous sinus lateral compartment (anatomical cadaver study). *Turk. Neurosurg.* **31**(5), 813–819 (2021).
35. Ceylan, S., Anik, I., Cabuk, B., Caklili, M. & Anik, Y. Extension pathways of pituitary adenomas with cavernous sinus involvement and its surgical approaches. *World Neurosurg.* **127**, e986–e995 (2019).
36. Yuanzhi, X. Intracranial breakthrough through cavernous sinus compartments: anatomic study and implications for pituitary adenoma surgery. *Oper. Neurol.* **23**(2), 115–124 (2022).

Acknowledgements
This work was supported by the Office of Ningbo Science and Technology (Grant Number 2012A610221). The authors also thank Prof. RS Dang, Department of Human Anatomy, Second Military Medical University, for assisting the material support and anatomic design.

Author contributions
Y.Y.: Project development, Data Collection, Manuscript writing; S.G. and H.H.: Data collection; G.Z.: Data Collection and analysis, Manuscript writing; J.L.: Project development; R.D.: Assisting the material support, Anatomic design. All authors reviewed the manuscript.

Competing interests
The authors declare no competing interests.

Additional information
Correspondence and requests for materials should be addressed to Y.Y.

Reprints and permissions information is available at www.nature.com/reprints.

Publisher’s note Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Open Access This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article’s Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article’s Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/.

© The Author(s) 2022