

Approaching Intradural Lesions of the Anterior Foramen Magnum and Craniocervical Junction: Anatomical Comparison of the Open Posterolateral and Anterior Extended Endonasal Endoscopic Approaches

This is the peer reviewed version of the following article:

Original:

Dannhoff, G., Gallinaro, P., Todeschi, J., Ganau, M., Spatola, G., Ollivier, I., et al. (2023). Approaching Intradural Lesions of the Anterior Foramen Magnum and Craniocervical Junction: Anatomical Comparison of the Open Posterolateral and Anterior Extended Endonasal Endoscopic Approaches. WORLD NEUROSURGERY, 178, e410-e420 [10.1016/j.wneu.2023.07.080].

Availability:

This version is available <http://hdl.handle.net/11365/1280810> since 2024-12-15T19:40:44Z

Published:

DOI: <http://doi.org/10.1016/j.wneu.2023.07.080>

Terms of use:

Open Access

The terms and conditions for the reuse of this version of the manuscript are specified in the publishing policy. Works made available under a Creative Commons license can be used according to the terms and conditions of said license.

For all terms of use and more information see the publisher's website.

(Article begins on next page)

Approaching intradural lesions of the anterior foramen magnum and cranio-cervical junction: anatomical comparison of the open postero-lateral and the anterior extended endonasal endoscopic approaches

Guillaume Dannhoff¹⁻³ MD, Paolo Gallinaro¹ MD*, Julien Todeschi¹ MD, Mario Ganau¹ MD-PhD, Giorgio Spatola¹ MD, Irène Ollivier¹ MD, Hélène Cebula¹ MD, Charles-Henry Mallereau¹, Seyyid Baloglu² MD, Raoul Pop² MD, Francois Proust¹ MD-PhD, Salvatore Chibbaro¹⁻³ MD-PhD.

*The first 2 authors have equally contributed to the study

¹ Department of Neurosurgery, Strasbourg University Hospital, Strasbourg France

² Neuroradiology Unit, Strasbourg University Hospital, Strasbourg France

³ Skull Base ENT-Neurosurgery Laboratory of the University Hospital of Strasbourg, France

Key words: Chordoma, Cranial base approaches; Far Lateral Approach; Far Medial Endonasal Approach; Foramen Magnum; Meningioma; Skull base surgery.

Short title: Anatomical comparison of anterior and postero-lateral approaches to the foramen magnum

Corresponding Author:

Dr. Guillaume Dannhoff MD
Strasbourg University Hospital
1, Av. Moliere, 67098 – Strasbourg France
guillaume.dannhoff@neurochirurgie.fr
guillaume.dannhoff@chru-strasbourg.fr

1 **ABSTRACT**

2 **Background:** Lesions of the foramen magnum (FM) and cranio-cervical junction (CCJ) area are traditionally managed
3 surgically through anterior, antero-lateral and postero-lateral skull-base approaches. This anatomical study aimed to
4 compare the usefulness of a modified extended endoscopic approach (EEA), the so-called far-medial endonasal
5 approach (FMEA), versus the traditional posterolateral far-lateral approach (FLA).

6 **Material and Methods:** Ten fixed silicon injected heads specimens were used in the Skull Base ENT-Neurosurgery
7 Laboratory of the University Hospital of Strasbourg, France. A total of 20 FLA and 10 FMEA were realized. A high-
8 resolution CT scan was performed for a quantitative analysis of the different approaches. The analysis aimed to estimate
9 the extent of surgical exposure and the freedom of movement (manoeuvrability) through the operating channel, using a
10 polygonal surface model to get a morphometric estimation of the area of interest (surface and volume) on post-
11 dissection CT scan using Slicer 3D software.

12 **Results:** FMEA allows a more direct route to the anterior FM, with a wider brainstem exposure compared to the FLA,
13 and an excellent visualization of all anterior midline structures. Limitations of FMEA include the deep and narrow
14 surgical corridor, and the difficulty to reach lesions located laterally over the jugular foramen and the hypoglossal canal.

15 **Conclusion:** FMEA and FLA are both effective surgical routes to reach FM and CCJ lesions; a good command of both
16 should be enrooted in any modern skull base surgeon as they appear complementary. This anatomical study provides
17 tools to comprehensive preoperative evaluation and selection of the most appropriate surgical approach.

18

19

20

21

22

23 INTRODUCTION

24 The occipital bone surrounding the foramen magnum is made of three portions, the basilar part or clivus anteriorly, the
25 squamous part posteriorly and the condylar part in-between, protruding inferiorly to form the atlanto-occipital joint.
26 Together, these portions encompass various neurovascular structures in a complex anatomical display ([Fig. 1](#)). Surgical
27 areas surrounding the foramen magnum, i.e. retro-clival area and cranio-cervical junction (CCJ) can be affected by
28 various intradural and extradural disorders, including: tumours, bone malformations, inflammatory diseases and trauma
29 ¹. Given that, several approaches to those regions have been developed through anterior, antero-lateral and postero-
30 lateral routes ²⁻⁴.

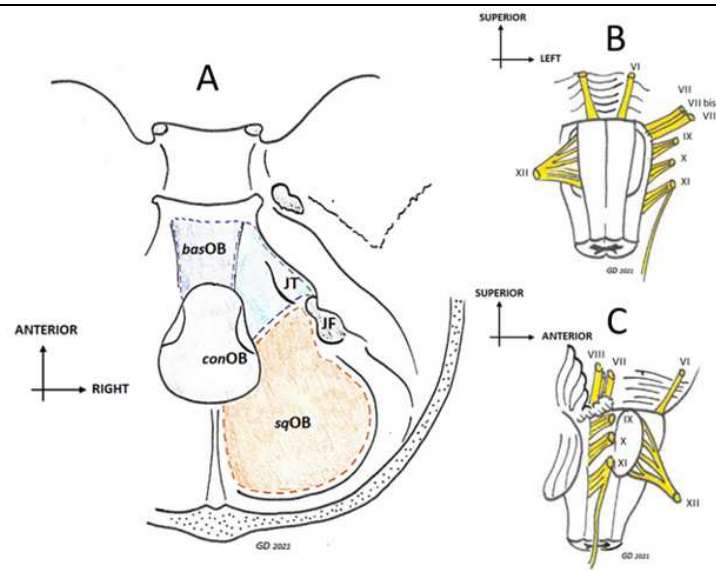
31 This variety of approaches reflects the difficulty to safely reach these regions. Neurosurgical corridors aim to provide
32 the best possible exposure of the surgical field with the minimal manipulation or retraction of anatomical structures and
33 therefore with limited morbidity. Nevertheless, many of these approaches are quite invasive, requiring a variable degree
34 of neurovascular manipulation.

35 The far-lateral approach (FLA) is an extension of the classical suboccipital approach, consisting in a lateral extension of
36 the craniotomy to reach the condylar fossa along with removal of the posterior arch of C1. Originally described in 1986
37 by Heros for the management of vertebrobasilar artery lesions ⁵, its popularity increased two years later with Bernard
38 George series of 14 benign anterior FM tumours successfully resected ⁶. It has since become a traditional way to access
39 lesions of the inferior clivus from a posterolateral perspective,

40 However, the most physiological route to the retro-clival area and the ventral CCJ would in theory be represented by
41 anterior approaches, offering a direct view of deep surgical targets through a corridor that is not crossed by cranial
42 nerves. Since its inception, the trans-nasal approach has been mainly used to treat extradural lesions and to perform CCJ
43 decompression ⁷⁻⁹, although some authors also have reported its use to manage intradural lesions ¹⁰. The recent
44 development of endoscopic endonasal approaches for the treatment of sellar region lesions ¹¹ has led some surgeons to
45 conceive the extended endonasal endoscopic approach (EEA), which is primarily aimed at reaching pathologies located
46 in suprasellar, parasellar, but also retroclival areas ¹²⁻¹⁶. Recent anatomical studies and clinical reports have also
47 demonstrated the possibility of directly approaching any midline structure, from the crista galli to the odontoid process,
48 by different EEAs ^{17,18}. Finally, the advancement of surgical technique with an improvement of neuro-endoscopic
49 technology led to an evolution of the EEA, including the so-called far-medial endonasal approach (FMEA), with
50 potential to treat lesions located in the inferior clival region (mainly extradural chordomas), as well as in the CCJ.

51 The FMEA was described by Morera in 2010 ¹⁹, consisting in an anterior endoscopic trans-clival approach optimizing
 52 surgical exposure via drilling of the lateral inferior clival area, together with a condylar and jugular tubercle partial
 53 resection. Endoscopic identification of the supracondylar groove is a key landmark to locate the hypoglossal canal,
 54 itself separating the jugular tubercle superiorly and the condylar compartment inferiorly.

55 The aim of the present study was to perform a morphometric analysis of a) the surgical exposure and b) the
 56 manoeuvrability through the operating channels obtained through the FLA and the FMEA (Fig. 1), and to eventually
 57 compare our results with those recently published in the neurosurgical literature.



58
 59 **Figure 1.** Schematic drawings of main osseous and nervous landmarks of surgical approaches to the foramen magnum. (A) Skull
 60 base drawing showing the bone drilling areas around the foramen magnum. The occipital bone is made of three portions, the basilar
 61 part (*basOB*) or clivus anteriorly, the squamous part (*sqOB*) posteriorly and the condylar part (*conOB*) in-between. The jugular
 62 foramen (JF) and jugular tubercle (JT) are two important lateral landmarks guiding surgical approaches. The standard endoscopic
 63 endonasal approach is represented by the dark blue dotted line, the extended FMEA by the light blue line and the classic FLA by the
 64 orange line respectively. (B, C) Schematic drawings of the brainstem exposed through surgical approaches to the foramen magnum:
 65 anterior view of the ventromedial brainstem approached via the FMEA (B) and right lateral view of the dorsolateral brainstem
 66 approached via the FLA (C).

69 MATERIAL AND METHODS

70 Dissections were performed on 10 fixed coloured silicone-injected adult head specimens, without previous brain or
71 skull base abnormalities, at the Skull Base ENT-Neurosurgery Laboratory of the University Hospital of Strasbourg,
72 France. Each specimen underwent a double FLA (for a total of 20 approaches) and a single FMEA (for a total of 10
73 approaches).

74 Morphometric Analysis

75 Before dissection, each specimen underwent a high-resolution computed tomography (CT) scan (1 mm cuts) to allow
76 accurate measurements of deep anatomical structures exposed during dissection along the surgical corridor. Though
77 MRI is the preferred imaging modality in clinical routine when dealing with FM lesions, CT scan was performed here
78 due to its availability in our research laboratory and also due to its high-quality bony definition and analysis, allowing
79 more precise and reproducible evaluation of our parameters of interest.

80 A post-dissection high-resolution CT scan was also obtained for the purpose of quantitative analysis of the different
81 approaches. The analysis was done using a polygonal surface model to get a morphometric estimation of the area of
82 interest (its surface and volume) on post-dissection CT scan ([Fig. 2, 3](#)); those calculations were conducted through the
83 Slicer 3D software (Version 4.10.1, www.slicer.org), which analysed 2 main features:

- 84 • extent of surgical exposure (surface and volume of exposure): considered as the maximal 2D/3D area that can
85 be exposed using each approach,
- 86 • the freedom of movement (surgical freedom): allowing to reach a specific target area without
87 retracting/damaging important neurovascular structures through the operating channel.

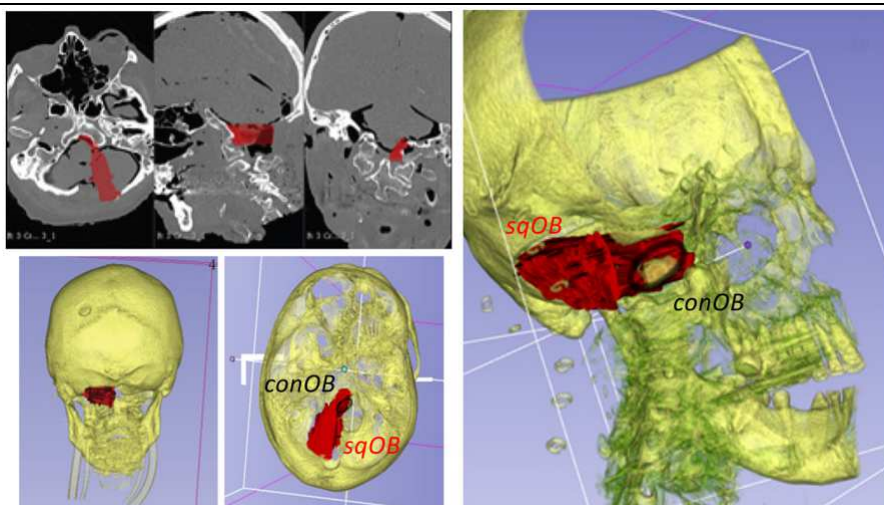


Fig. 2. CT-scan with 3D reconstructions of the surgical volume involved in FLA, accessing the foramen magnum posteriorly through resection of the squamous portion of the occipital bone (*sqOB*) and partial resection of the occipital condyle (*conOB*) highlighted in red.

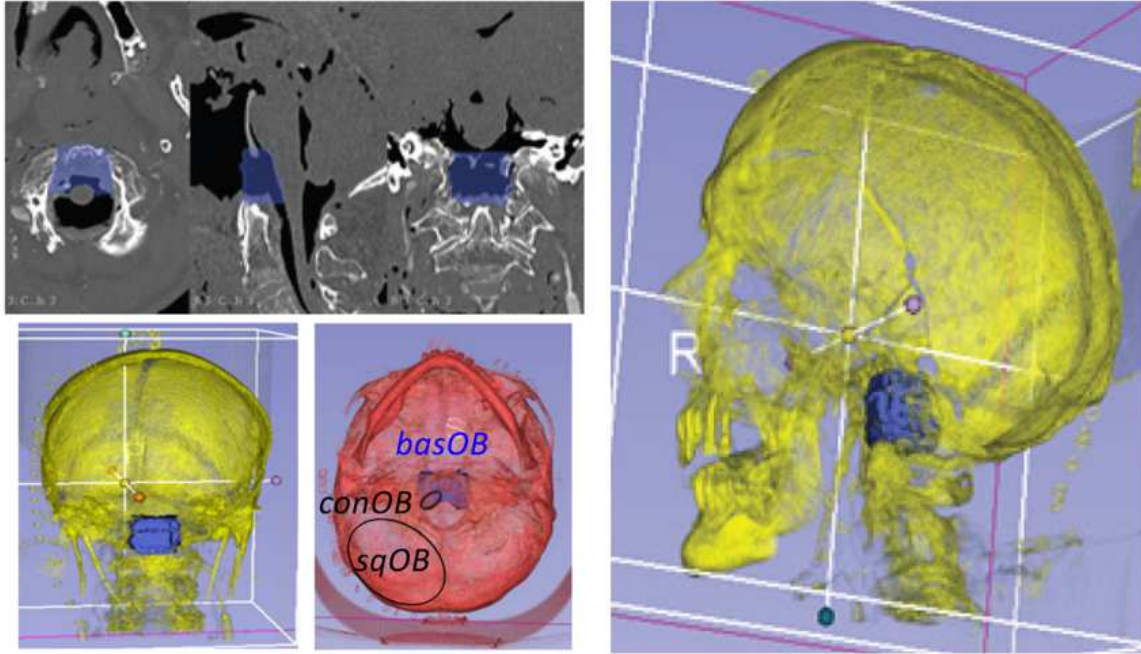


Fig. 3. CT-scan with 3D reconstructions of the surgical volume involving the FMEA, accessing the foramen magnum anteriorly through resection of the basilar portion of the occipital bone (*basOB*) highlighted in dark blue.

Step-by-step description of surgical technique

The FLA was performed bilaterally in all specimens, under microscopic magnification (Zeiss surgical microscope), with the heads fixed in a three-pin Mayfield head holder, in a surgical position (i.e. prone position and slightly rotated 15° to the opposite side). Dissection was performed according to the techniques previously described in the literature, consisting of three critical steps (Fig. 4) :

1°) A “hockey stick” skin incision is performed, extending from the spinous process of C3 to the inion, then curving laterally toward the apex of the mastoid process. Muscles are dissected from the occipital bone and posterior arch of C1, and retracted laterally. At the end of this first step, we identify the suboccipital triangle and the vertebral artery in its V3 segment.

2°) A C1 hemilaminectomy is done, extending from a point just beyond the midline to the groove of the vertebral artery. The vertebral artery is not transposed, to simulate the common usage of FLA in clinical practice. Once the

107 posterior arch of C1 is removed, the posterior atlantooccipital membrane can be detached superiorly from the posterior
 108 margin of the FM, allowing then to perform a lateral suboccipital craniotomy. The craniotomy involves the squamous
 109 portion of the occipital bone, extending laterally until being a few millimeter away from the neurovascular components
 110 of the jugular foramen. Bone resection is in fine limited by venous structures – namely the sigmoid sinus laterally and
 111 the transverse sinus superiorly -. Finally, the exposed occipital condyle is minimally drilled to reproduce a
 112 transcondylar variant of the FLA. In order to preserve the stability of the cranio-cervical junction and to ensure the
 113 protection of the vertebral artery, condylectomy is limited to its medial third, bordered anteriorly by the hypoglossal
 114 canal and inferiorly by the articular facet of the lateral mass of C1.



115
 116 **Fig. 4** Dissection steps of the FLA, from hockey stick incision to C1 hemilaminectomy and occipital craniotomy with
 117 partial condylectomy allowing wide dural exposure.

118 3°) The dura mater is finally opened in the midline, from the height of the dural entry point of the vertebral artery, and
 119 curved to the sigmoid sinus margin cranially. The cerebellar hemisphere is then gently retracted until reaching the
 120 inferior cranial nerves, vertebral artery and mostly until adequate exposure of the inferior clivus is achieved.

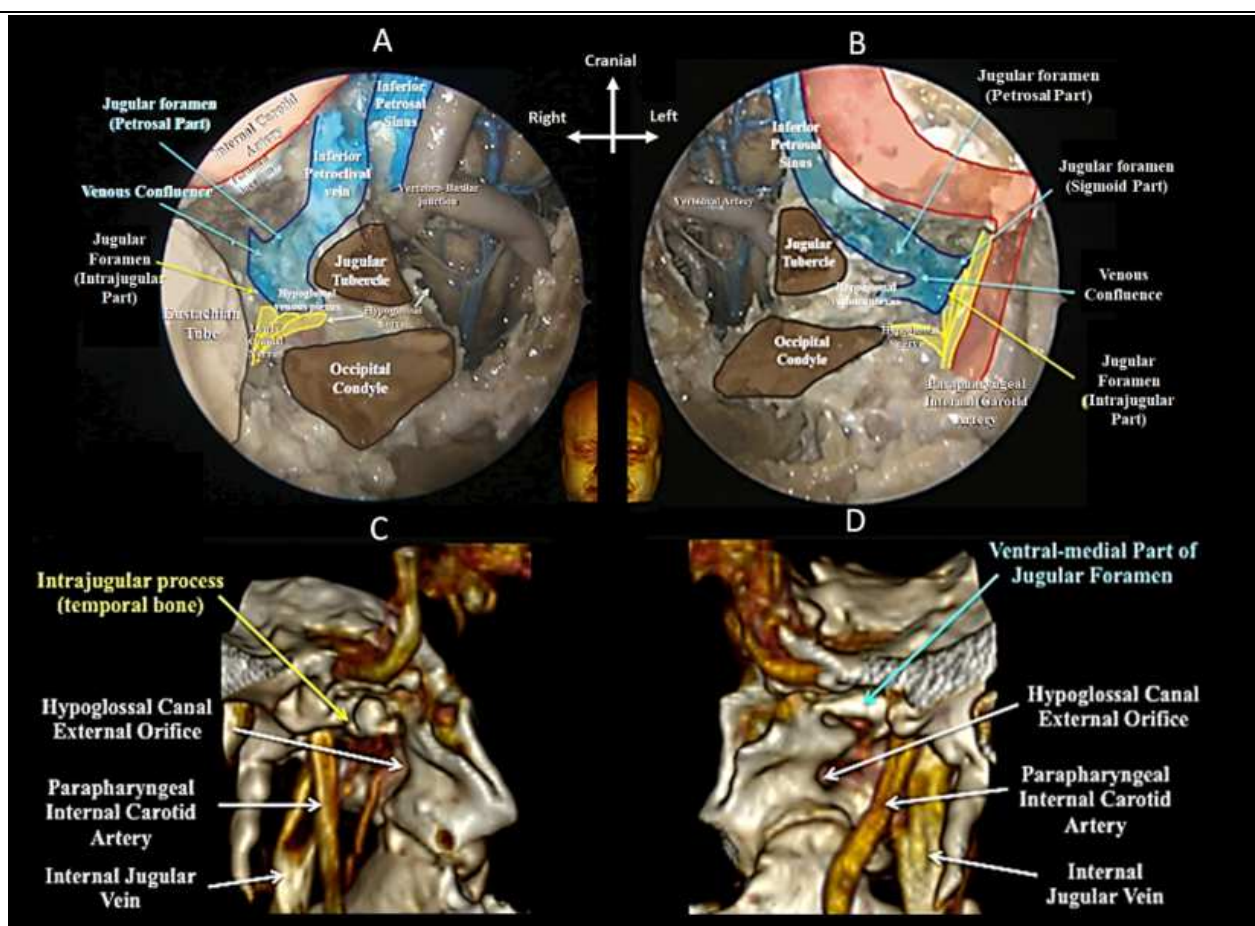
121 The FMEA is performed with a 4 mm diameter, 18 cm long endoscope with 0° and 30 and sometimes 45° optics, and a
 122 TELEPACK X LED image system (Karl Storz GmbH & Co., Tuttlingen, Germany), according to the technique
 123 previously described in the literature in the way to expose the FM and inferior clivus along with the condylar region,
 124 jugular foramen, jugular tubercle and cranio-cervical junction. The heads were fixed in a Mayfield three-pin headholder
 125 and positioned with approximately 15° flexion and 10° rotation to the right, to reproduce the common patient setup in
 126 the operating room.

127 The FMEA is performed via a bilateral transnasal approach, through the following steps:

128 1°) A bilateral inferior turbinectomy is performed to enlarge the surgical corridor, while taking care of preserving the
 129 nasal mucosa flap bilaterally. The mucosal and muscular planes, made by the posterior wall of the nasopharynx, are

130 resected to expose the inferior clivus and the cranial aspect of the craniocervical junction. The level of the jugular
 131 foramen and the hypoglossal canal are localized through the identification of the pharyngeal tubercle and the
 132 supracondylar groove respectively.

133 2°) The atlanto-occipital joint capsule is removed to expose the occipital condyle. The clivus is drilled off from the
 134 pharyngeal tubercle to the foramen magnum. The superolateral limit of the clivectomy is defined by the lateral
 135 pharyngeal tubercle, as going beyond it could result in damaging the internal carotid artery superolaterally or the
 136 inferior petrosal sinus and jugular bulb laterally. The ventral aspect of the condyle is drilled up to the cortical bone of
 137 the hypoglossal canal, which constitutes the lateral limit of the clival bony removal (Fig. 5).



138
 139 **Fig. 5.** Endoscopic view of the FM region approached anteriorly after clival resection, showing the ventral brainstem
 140 along with the variety of osseous landmarks and neurovascular structures visualized through a narrow surgical corridor
 141 on the right (A) and left (B) sides. The clival drilling area is limited by the hypoglossal canal above the occipital
 142 condyle and the jugular tubercle viewed bilaterally. Per-procedural findings are correlated with 3D reconstructions of
 143 the corresponding CT-scans right (C) and left (D) aspects.

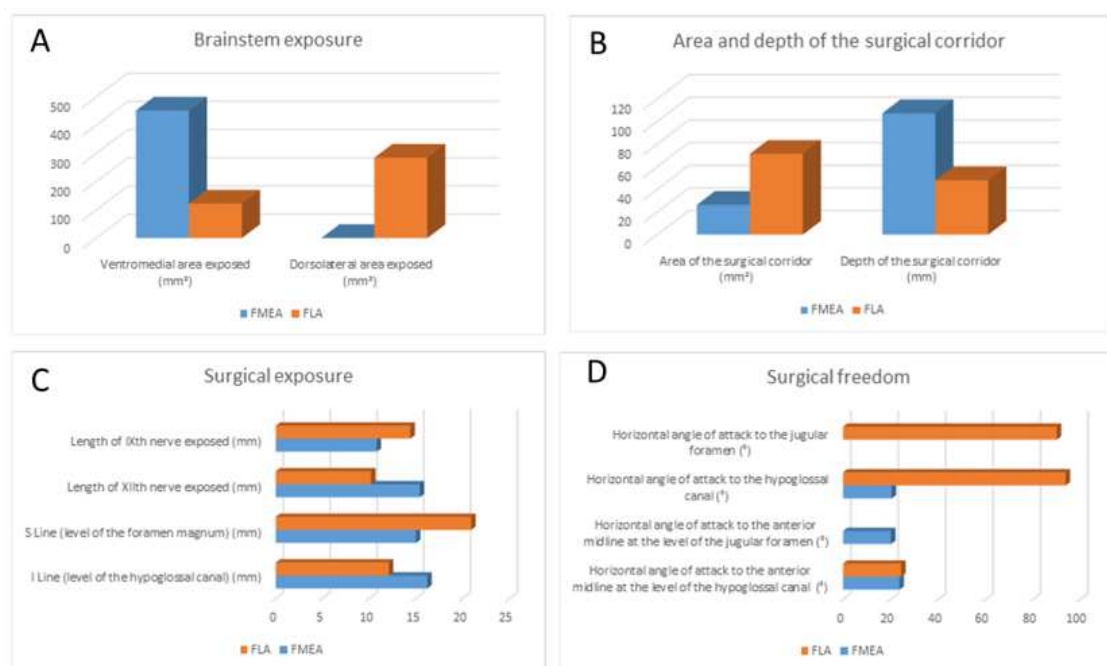
145 3°) Finally the dura mater is opened in the midline, allowing visualisation of the brainstem, the vertebral, basilar and
 146 origin of the posteroinferior cerebellar arteries, along with the cranial nerves IX, X, XI and XII bilaterally. (Fig. 5)

147 Surgical exposure was quantified during dissections using a paper graduated ruler and was correlated with post-
 148 dissection measures based on CT-scan.

149 **Statistical analysis**

150 A paired Student t-tests was used to compare the data for quantitative analysis. In the statistical comparison analysis, a p
 151 value < 0.05 was considered significant. All calculations were performed with GraphPad Prism version 6.00 (GraphPad,
 152 La Jolla, California, USA)

153 **RESULTS**



154
 155 **Fig. 6.** Main parameters quantified when performing FLA (orange bars) and FMEA (blue bars): (A) Brainstem
 156 exposure (B) Area and depth of the surgical corridor provided (C) Surgical exposure of cranial nerves IX and XII, and
 157 reference lines respectively defined at the level of the foramen magnum and at the level of the hypoglossal canal. (D)
 158 Surgical freedom obtained at the level of interest.

159

160

161

162 **Brainstem exposure**

163 FMEA allowed a significantly wider exposure ($p < 0.01$) of the ventral brainstem, up to 180° of its ventro-medial part
164 ($452.4 \pm 14.33 \text{ mm}^2$), compared to the FLA ($122.4 \pm 4.58 \text{ mm}^2$) ([Fig. 6A](#)). In practice, through the FMEA it was
165 possible to obtain a total exposure of the ventral brainstem, whereas only 25% was exposed with the unilateral FLA.

166 The widest exposure obtained with the FLA was $407.6 \pm 15.89 \text{ mm}^2$ (40% at the level of the ventro-medial area and
167 60% at the level of the dorso-lateral area). More specifically, the dorso-lateral area of the brainstem exposed through the
168 FLA was of $285.2 \pm 11.31 \text{ mm}^2$, corresponding to 75% of the entire dorso-lateral face of the brainstem. On the other
169 hand, through the FMEA, it was not possible to expose any aspect of the dorsolateral area of the brainstem, because of
170 the cranial nerves crossing over.

171 **Surgical corridor**

172 The FLA created a surgical corridor up to the ventro-medial area of the posterior fossa in only 70% of the specimens; in
173 the remaining 30% (6/20) the access to the clivus was limited by the presence of a homolateral predominant vertebral
174 artery ([Fig. 6B](#)). The mean sectional area of the surgical corridor in the FLA was about $71.4 \pm 3.86 \text{ mm}^2$, whereas the
175 mean sectional area of the FMEA was about $26.4 \pm 3.65 \text{ mm}^2$. The possibility to gently retract the cerebellar
176 hemisphere, together with the wider surgical angle offered by the FLA resulted in a wider surgical corridor than that
177 offered by FMEA. Of note, in 40% of the specimens the spinal root of the XI cranial nerve divided the main surgical
178 corridor created by FLA into two corridors, superior and inferior, each with similar dimensions.

179 In FLA the depth of the surgical corridor corresponded to the distance between the musculocutaneous flap and the dura
180 mater, its length was on average $48.0 \pm 2.74 \text{ mm}$; on the other hand, in FMEA the depth of the surgical corridor
181 corresponded to the distance between the anterior nasal spine and the dura mater, its average length was 107.2 ± 3.70
182 mm.

183 **Surgical exposure**

184 Two key lines were calculated in all specimens: the S Line, or the average distance that could be reached in the axial
185 plane at the level of the foramen magnum, and the I line, or the average distance that could be reached in the axial plane
186 at the level of the hypoglossal canal. The S Line was calculated in the latero-medial sense to the midline for FLA and in
187 the medio-lateral sense to the periphery for FMEA and resulted significantly higher in FLA than in FMEA: $20.9 \pm$
188 1.20 mm versus $15.0 \pm 0.72 \text{ mm}$, $p < 0,01$ ([Fig. 6C](#)). The I line was calculated in the latero-medial sense for FLA and
189 in the medio-lateral sense for FMEA and resulted significantly higher in FMEA than in FLA: $16,2 \pm 0,84 \text{ mm}$ versus
190 $12,1 \pm 0,99 \text{ mm}$, $p < 0,01$.

191 Of note, the average intradural length of the IX cranial nerve exposed by FLA was significantly longer than the
192 exposure provided by FMEA: 14,4 +/- 1,37 mm versus 10.8 +/- 1.30 mm, $p < 0.05$. On the contrary, the average
193 intradural length of the XII cranial nerve exposed by FMEA was significantly longer than that offered by FLA: 15.4 +/-
194 1.14 mm versus 10.2 +/- 1.03 mm, $p < 0.01$.

195 **Surgical freedom**

196 The extent of surgical exposure and the freedom of movement to the jugular foramen could only be assessed in the
197 corridor created through a FLA, its average value was $90.1^\circ \pm 6.61^\circ$ ([Fig. 6D](#)). In our dissected specimen, the jugular
198 foramen was indeed not reachable at all through a FMEA. On average, the horizontal angle of attack to the hypoglossal
199 canal was significantly higher in FLA than in FMEA: $93.9^\circ \pm 5.51$ versus $20.5^\circ \pm 1.05$, $p < 0.01$.

200 The anterior midline at the level of the jugular foramen could not be reached in any anatomical subject through a FLA.
201 Though it was accessible in every subject through FMEA, its angle of attack to the anterior midline was limited (20.2°
202 ± 1.48). The anterior midline at the level of the hypoglossal canal was accessible in 40% of cases via FLA and 100%
203 of cases via FMEA, with an almost identical angle of attack, ranging between 23.8° and 24.5° in both types of
204 approach.

205

206 **DISCUSSION**

207 The FM is the largest foramen in the occipital bone of the skull in humans and many other animals. The spinal cord, an
208 extension of the medulla oblongata, passes through the FM as it exits the cranial cavity. Apart from the medulla
209 oblongata, also the vertebral arteries, the anterior and posterior spinal arteries, the tectorial membranes, alar ligaments,
210 and the accessory nerve transit through the FM. This foramen extends anteriorly from the junction of the lower and
211 middle third of the clivus to the upper edge of the C2 body, laterally from the jugular tubercle to the upper aspect of the
212 C2 lamina, and posteriorly from the anterior edge of the occipital bone to the C2 spinous process^{17,18}.

213 **Dissection Results**

214 Access to the anterior FM and to the CCJ remains one of the most complex neurosurgical procedures because of the
215 proximity to vital neurovascular structures, depth and narrowness of surgical corridors¹. In the literature, several
216 surgical approaches have been described for this region, each one with its pros and cons^{8,20}. In recent years thanks to
217 the progresses in terms of anatomical knowledge of the skull base and the development of endoscopic techniques and
218 the dedicated instrumentation, a new opportunities emerged to exploit the natural endonasal corridor and reach the
219 lower clivus and the CCJ^{2,18,21}.

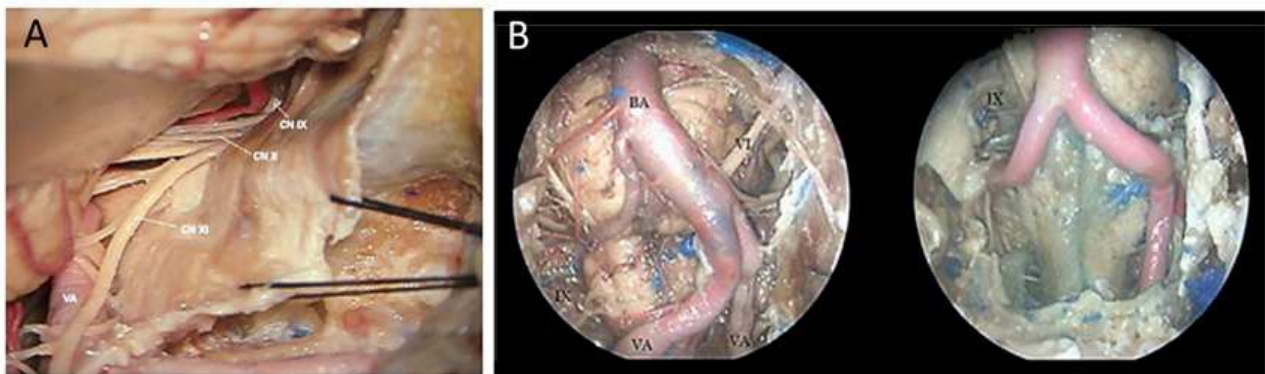
220 Having gathered as a group a 30-year experience with the FLA and a 15-year experience with the EEA ^{4,16}, we designed
221 an anatomical study to compare the extent of surgical exposure and the freedom of movement (manoeuvrability)
222 through both operating channels, FLA and FMEA.

223 The FLA is very popular worldwide due to its lower rate of complications compared to the extreme-lateral approach, it
224 provides a safe access to the anterior midline, although increasing the risk of vascular complications ^{3,20,22}.

225 Most results from our anatomical study are in line with previous similar studies ^{20,22}, this is not surprising given the
226 homogeneity of methodology for what regard morphometric analyses (we performed similar types of calculations with
227 the same software). This allowed our data to be comparable to those obtained from previous studies, and increase the
228 generalizability of the conclusions reached so far by other authors.

229 Our results confirm that the surgical exposure of the ventral surface of the FM and of the inferior clivus is far superior
230 via FMEA; however the manoeuvrability is lower in this latter technique because of the narrow endonasal corridor.

231 **“Open” FLA**



232
233 **Fig. 7** Dissection views comparing neurovascular structures exposed via the FLA (A) and the FMEA (B) after dural
234 opening. BA: basilar artery; VA: vertebral artery; VI : sixth cranial nerve; IX: ninth cranial nerve;

235 The FLA is widely used in the treatment of FM lesions, but is also utilized for anterolateral lesions of the inferior clivus,
236 because it allows an easy occipital condyle drilling, with the possibility to release and displace the vertebral artery (VA)
237 and to drill the jugular tuberculum ^{9,23,24}. In fact the possibility to expose and control the VA in the early stage of the
238 procedure allows to reduce the risk of bleeding during the resection of an inferior clivus lesion, and represents another
239 advantage of the FLA.

240 In our study, as previously reported, the FLA allowed a good exposure of the inferior clivus as well as of the
241 anterolateral side of the FM and the ventrolateral surface of the brainstem together with the related neurovascular
242 structures, all without the need of traction ^{20,22,25} ([Fig. 7A](#)). This approach also allows a wide exposure of the dorso-

lateral area of the posterior cranial fossa²⁰. It should be pointed out, however, that the FLA offers a very limited access to the ventromedial brainstem area, in particular to the anterior midline at the cranial level. In fact, in our study it was possible to reach the clivus in only 30% of specimens due to the anatomical obstruction caused by a dominant VA; whereas at the level of the hypoglossal canal, the anterior midline could be reached in only 40% of specimens. Of note, at the level of the jugular foramen, the anterior midline could never be reached. Because of these limitations, the distance reached in a medial direction to the inferior clivus was of 21 mm at the level of the jugular foramen and of 12 mm at the level of the hypoglossal canal.

FMEA

Since its introduction, EEA allowed to overcome several limitations of conventional microsurgical approaches. For instance, it allowed a panoramic and dynamic vision in a narrow surgical corridor with visualisation of numerous landmarks, while guaranteeing excellent image magnification when exploring neurovascular structures ([Fig. 7B](#)). With time, neurosurgeons started to use such approach to tackle lesions of the anterior and posterior cranial fossa^{17-19,26,27}.

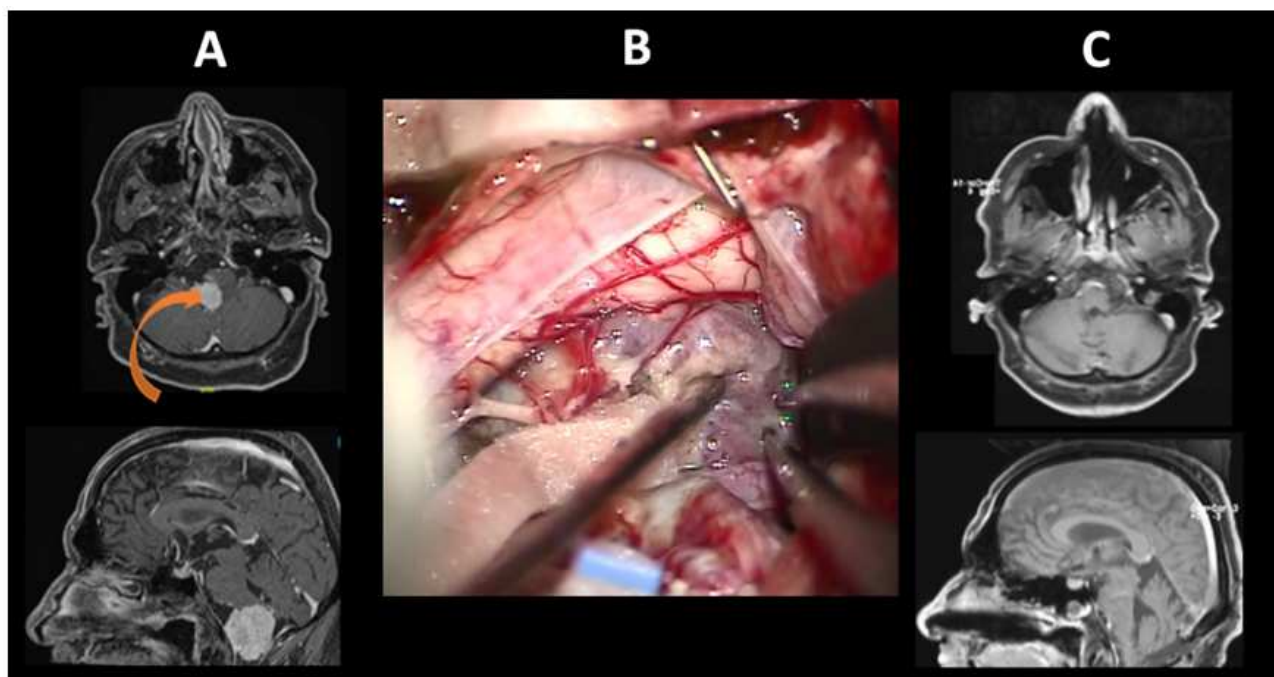
In our study, through FMEA the depth of the surgical corridor to the FM had an average of 10.5 cm and was significantly higher to that to the CCJ provided by FLA: because of the narrow surgical corridor, the FMEA is characterized by a lower grade of surgical freedom, compared to the open surgical approaches, in particular for what concern the laterality. Despite these limitations, our results showed that the FMEA allows an excellent exposure and a direct access to the medial part of the anterior FM and to the inferior clivus, with an average distance of 1.5 cm from the midline of the jugular foramen, and of 1.8 cm from the hypoglossal canal. However, the lateral exposure of the lesion is limited by some uncrossable borders such as internal carotid arteries (ICAs), nerve of the pterygoid canal (vidian nerve), jugular veins, hypoglossal nerve and Eustachian tubes²⁸. The degree of surgical freedom is also limited by the handling of the endoscope, because of anatomical obstacles such as the nose, the nasal concha and the depth of the endonasal corridor itself. Because of the above described limitations, the FMEA does not allow, in our opinion, a good exposure of lesions localized in the antero-lateral region of the FM and of the inferior clivus, in particular for those whose epicentre is localized at the level of the jugular foramen or at the hypoglossal canal. In fairness, the FMEA should not be considered as a mini-invasive approach because it is really destructive and can lead to important complications, such as the postoperative discomfort related to the turbinectomy, the risk of instability at the level of the CCJ and, most importantly, the risk of meningitis and CSF leak in case of intradural lesions²⁹. According to our results, the FMEA is particularly useful for lesions with a small volume and near to the anterior midline (no further than 1.5 cm from the midline).

272 In order to allow a safe tumour resection through the endoscope, the caudal extension of the lesion should be limited: all
273 tumours that cross the odontoid process are more difficult to treat because of the conflict between the endoscopic
274 instrumentations, the anterior nasal spine and the palatal plate. At last, the extent of the resection over the odontoid
275 process could compromise the stability of the CCJ by damaging the anterior longitudinal ligament ²⁷. As it has been
276 previously showed by Benet et al., the FMEA could be also considered in a selected group of patients (characterized by
277 good general health, potential healing after a gross total resection of the lesion and tumours with extensions in the
278 ventro-medial and dorso-lateral areas) for combined open FLA in order to achieve a gross total resection of the lesion.

279 Clinical applications examples

280 Case vignette 1

281 A 56-year-old woman presented with a large clival meningioma, centred on the right anterolateral portion of the FM.
282 The vertebral arteries were not encased, the right one being located rather anteriorly and the left one laterally; the brain
283 stem was severely stretched and displaced postero-laterally to the left (Fig. 8A). The Far-Lateral Approach was judged
284 particularly adequate as it exploited the anatomical distortions and the surgical corridor created by the lesion itself (Fig.
285 8B). Moreover, section of the denticulate ligament allowed gentle mobilization of the upper spinal cord, increasing
286 surgical manoeuvrability. The patient was operated in a prone position, trough the traditional aforementioned steps
287 achieving a complete excision (Fig. 8C).



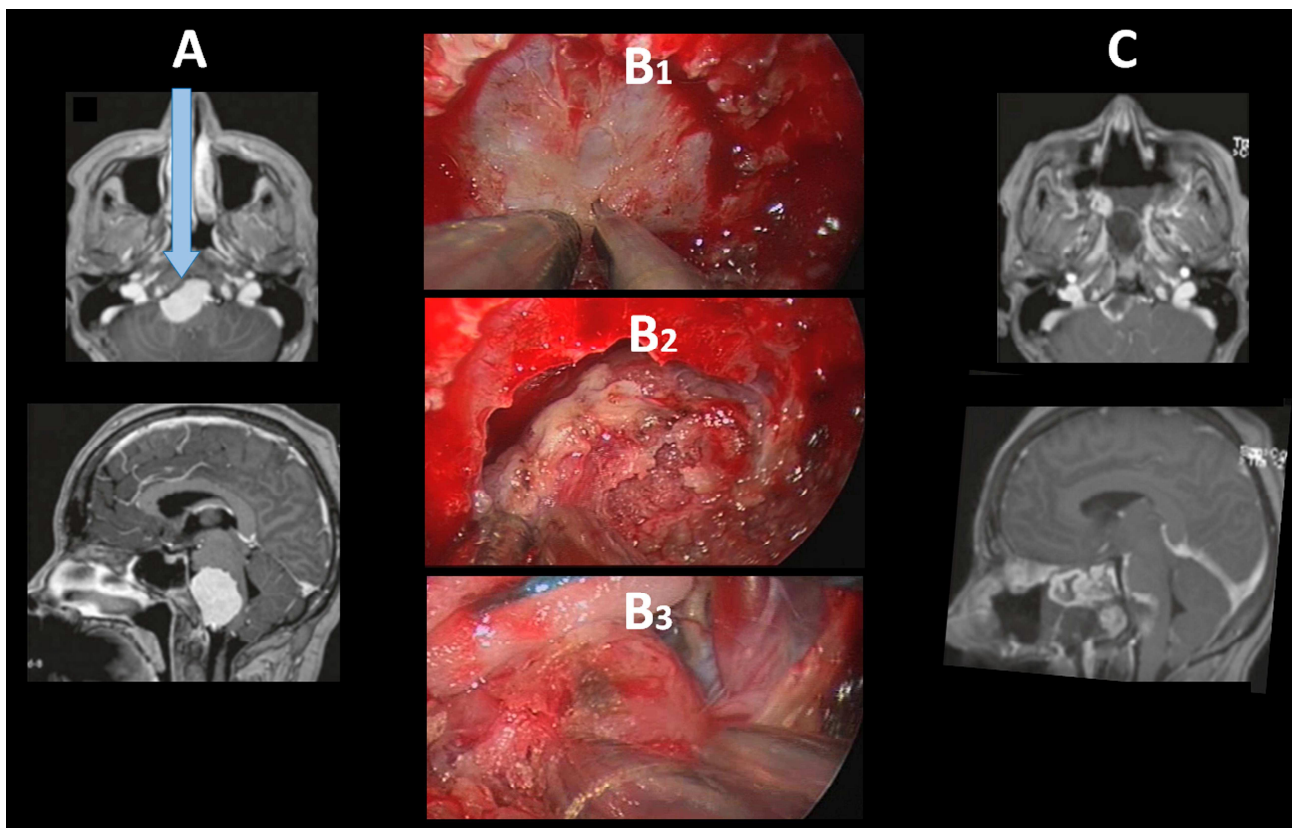
288
289 **Fig. 8** Case vignette: FLA resection of a clival meningioma, preoperative MRI (A), perioperative view of FLA (B) and
290 postoperative MRI showing complete removal (C).

291

292

293 Case vignette 2

294 A 47-year old woman presented with a FM meningioma. The epicentre of the lesion was in the ventromedial
 295 compartment; it did not extend beyond 1.5 cm from the anterior midline and did not overpass the jugular foramen and
 296 the hypoglossal canal laterally (Fig. 9A). The cranial nerves and vertebral artery were located postero-laterally. The
 297 caudal extension of the lesion did not exceed the apex of the odontoid. In this case, an Endo-FM approach was chosen
 298 (Fig. 9B), allowing adequate exposure so that complete excision could be achieved (Fig. 9C).



299

300 **Fig. 9** Case vignette: FMEA resection of a clival meningioma, preoperative MRI (A), peroperative view allowed with
 301 FMEA (B) showing dural exposure (B₁) followed by tumour exposure (B₂) and visualization of leptomeninges and
 302 neurovascular structures along tumour resection (B₃), postoperative MRI showing complete removal (C).

303 Study limitations and strengths

304 Our study is affected by a number of limitations. First of all, the small number of specimens studied: only 10 heads for a
 305 total of 20 FLA and 10 FMEA; as a consequence, the morphometric data obtained could have limited significance and

306 should be improved by further studies. Another limitation is represented by the specimens themselves. Because of the
307 fixation, there was a higher level of tissue rigidity, with a lower possibility of traction on neurovascular structures.

308 At last, it should be specified that the heads used for the dissection were not characterized by pathologies in the
309 anatomical area of interest. Understanding of normal anatomy is necessary to apprehension of disrupted anatomy. But
310 extrapolation of our anatomical data to pathological scenarios are limited, as the normal anatomy here depicted could be
311 variously disrupted by lesions' mass effect, with creation of natural surgical corridors and cleavage planes that need to
312 be assessed on a case-by-case basis.

313 This anatomical study could therefore benefit from more robust clinical perspectives. We firmly believe that when
314 enhanced with the experience of real-life surgical experience, those anatomical data could lead to the proposition of a
315 clinical algorithm allowing to select the most appropriate surgical approach.

316

317 **CONCLUSION**

318 Data from this anatomical study align well with other studies in the literature and demonstrate that FMEA and FLA
319 complement each other in the way both provide valuable treatment options for the surgical management of FM and CCJ
320 lesions. A careful pre-surgical planning under the “first do not harm” principle should guide neurosurgeons in opting for
321 an endoscopic or open surgical approach while choosing, on a case-by-case basis, the best technique to obtain the
322 maximal safe resection with minimal risk of iatrogenic complications. Our study suggests that rigorous anatomical
323 knowledge, together with a good surgical dexterity in those working channels, are the key to safely manage such
324 challenging lesions. The future endeavour for skull base surgeons then becomes the identification of patients and lesions
325 characteristics allowing for a comprehensive preoperative evaluation and the selection of the most appropriate surgical
326 approach, this further step should be based on objective evidence from nomograms and should spur into robust
327 operative algorithms.

328 **REFERENCES**

- 329 1. Samii M, Gerganov VM. Tumors of the occipitocervical junction. In: *Pathology and Surgery around the Vertebral*
330 *Artery*. Springer Paris; 2011:427-441. doi:10.1007/978-2-287-89787-0_29
- 331 2. Komotar R, Zacharia B, McGovern R, Sisti M, Bruce J, D'Ambrosio A. Approaches to anterior and anterolateral
332 foramen magnum lesions: A critical review. *J Craniovert Jun Spine*. 2010;1(2):86. doi:10.4103/0974-8237.77672
- 333 3. Leon-Ariza DS, Campero A, Romero Chaparro RJ, Prada DG, Vargas Grau G, Rhoton AL. Key Aspects in
334 Foramen Magnum Meningiomas: From Old Neuroanatomical Conceptions to Current Far Lateral Neurosurgical
335 Intervention. *World Neurosurgery*. 2017;106:477-483. doi:10.1016/j.wneu.2017.07.029

- 336 4. Chibbaro S, Cornelius JF, Froelich S, et al. Endoscopic endonasal approach in the management of skull base
337 chordomas—clinical experience on a large series, technique, outcome, and pitfalls. *Neurosurg Rev.*
338 2014;37(2):217-225. doi:10.1007/s10143-013-0503-9
- 339 5. Heros RC. Lateral suboccipital approach for vertebral and vertebrobasilar artery lesions. *Journal of Neurosurgery.*
340 1986;64(4):559-562. doi:10.3171/jns.1986.64.4.0559
- 341 6. George B, Dematons C, Cophignon J. Lateral approach to the anterior portion of the foramen magnum. *Surgical*
342 *Neurology.* 1988;29(6):484-490. doi:10.1016/0090-3019(88)90145-0
- 343 7. Bruneau M, George B. Classification system of foramen magnum meningiomas. *J Craniovert Jun Spine.*
344 2010;1(1):10. doi:10.4103/0974-8237.65476
- 345 8. Doglietto F, Belotti F, Qiu J, et al. Endonasal and Transoral Approaches to the Craniovertebral Junction: A
346 Quantitative Anatomical Study. In: Visocchi M, ed. *New Trends in Craniovertebral Junction Surgery.* Vol 125.
347 Acta Neurochirurgica Supplement. Springer International Publishing; 2019:37-44. doi:10.1007/978-3-319-62515-
348 7_6
- 349 9. Matsushima T, Natori Y, Katsuta T, Ikezaki K, Fukui M, Rhoton AL. Microsurgical Anatomy for Lateral
350 Approaches to the Foramen Magnum with Special Reference to Transcondylar Fossa (Supracondylar Transjugular
351 Tubercle) Approach. *Skull Base.* 1998;8(03):119-125. doi:10.1055/s-2008-1058570
- 352 10. Talacchi A, Biroli A, Soda C, Masotto B, Bricolo A. Surgical management of ventral and ventrolateral foramen
353 magnum meningiomas: report on a 64-case series and review of the literature. *Neurosurg Rev.* 2012;35(3):359-368.
354 doi:10.1007/s10143-012-0381-6
- 355 11. Chibbaro S, Signorelli F, Milani D, et al. Primary Endoscopic Endonasal Management of Giant Pituitary
356 Adenomas: Outcome and Pitfalls from a Large Prospective Multicenter Experience. *Cancers.* 2021;13(14):3603.
357 doi:10.3390/cancers13143603
- 358 12. Neligan PC, Mulholland S, Irish J, et al. Flap Selection in Cranial Base Reconstruction: *Plastic and Reconstructive*
359 *Surgery.* 1996;98(7):1159-1166. doi:10.1097/00006534-199612000-00005
- 360 13. Fortes FSG, Carrau RL, Snyderman CH, et al. Transpterygoid Transposition of a Temporoparietal Fascia Flap: A
361 New Method for Skull Base Reconstruction after Endoscopic Expanded Endonasal Approaches. *The*
362 *Laryngoscope.* 2007;117(6):970-976. doi:10.1097/MLG.0b013e3180471482
- 363 14. La Corte E, Aldana PR, Ferroli P, et al. The rhinopalatine line as a reliable predictor of the inferior extent of
364 endonasal odontoidectomies. *FOC.* 2015;38(4):E16. doi:10.3171/2015.1.FOCUS14777
- 365 15. Carrabba G, Dehdashti AR, Gentili F. Surgery for clival lesions: open resection versus the expanded endoscopic
366 endonasal approach. *FOC.* 2008;25(6):E7. doi:10.3171/FOC.2008.25.12.E7
- 367 16. Chibbaro S, Ganau M, Cebula H, et al. The Endonasal Endoscopic Approach to Pathologies of the Anterior
368 Craniocervical Junction: Analytical Review of Cases Treated at Four European Neurosurgical Centres. In:
369 Visocchi M, ed. *New Trends in Craniovertebral Junction Surgery.* Vol 125. Acta Neurochirurgica Supplement.
370 Springer International Publishing; 2019:187-195. doi:10.1007/978-3-319-62515-7_28
- 371 17. Cavallo LM, Cappabianca P, Messina A, et al. The extended endoscopic endonasal approach to the clivus and
372 cranio-vertebral junction: anatomical study. *Childs Nerv Syst.* 2007;23(6):665-671. doi:10.1007/s00381-007-0332-
373 7
- 374 18. d'Avella E, Angileri F, de Notaris M, et al. Extended endoscopic endonasal transclival approach to the
375 ventrolateral brainstem and related cisternal spaces: anatomical study. *Neurosurg Rev.* 2014;37(2):253-260.
376 doi:10.1007/s10143-014-0526-x
- 377 19. Morera VA, Fernandez-Miranda JC, Prevedello DM, et al. “Far-Medial” Expanded Endonasal Approach to the
378 Inferior Third of the Clivus. *Operative Neurosurgery.* 2010;66(suppl_2):ons211-ons220.
379 doi:10.1227/01.NEU.0000369926.01891.5D

20. Benet A, Prevedello DM, Carrau RL, et al. Comparative Analysis of the Transcranial “Far Lateral” and Endoscopic Endonasal “Far Medial” Approaches: Surgical Anatomy and Clinical Illustration. *World Neurosurgery*. 2014;81(2):385-396. doi:10.1016/j.wneu.2013.01.091
21. Fang CH, Friedman R, Schild SD, et al. Purely endoscopic endonasal surgery of the craniovertebral junction: A systematic review: Endonasal craniovertebral junction surgery. *International Forum of Allergy and Rhinology*. 2015;5(8):754-760. doi:10.1002/alr.21537
22. Wang M, Chae R, Joseph Shehata, et al. Comparative Analysis of the Subtonsillar, Far-Lateral, Extreme-Lateral, and Endoscopic Far-Medial Approaches to the Lower Clivus: An Anatomical Cadaver Study. *World Neurosurgery*. 2019;127:e1083-e1096. doi:10.1016/j.wneu.2019.04.048
23. Di G, Fang X, Hu Q, Zhou W, Jiang X. A Microanatomical Study of the Far Lateral Approach. *World Neurosurgery*. 2019;127:e932-e942. doi:10.1016/j.wneu.2019.04.004
24. Vishteh AG, Crawford NR, Melton MS, Spetzler RF, Sonntag VKH, Dickman CA. Stability of the craniovertebral junction after unilateral occipital condyle resection: a biomechanical study. *Journal of Neurosurgery: Spine*. 1999;90(1):91-98. doi:10.3171/spi.1999.90.1.0091
25. Pai SB, Raghuram G, Keshav GC, Rodrigues E. Far-lateral Transcondylar Approach to Anterior Foramen Magnum Lesions - Our Experience. *Asian J Neurosurg*. 2018;13(3):651-655. doi:10.4103/ajns.AJNS_273_16
26. Singh H, Harrop J, Schiffmacher P, Rosen M, Evans J. Ventral Surgical Approaches to Craniovertebral Junction Chordomas. *Neurosurgery*. 2010;66(suppl_3):A96-A103. doi:10.1227/01.NEU.0000365855.12257.D1
27. Chibbaro S, Cebula H, Aldea S, et al. Endonasal Endoscopic Odontoidectomy in Ventral Diseases of the Craniocervical Junction: Results of a Multicenter Experience. *World Neurosurgery*. 2017;106:382-393. doi:10.1016/j.wneu.2017.06.148
28. Jho HD, Ha HG. Endoscopic endonasal skull base surgery: Part 3--The clivus and posterior fossa. *Minim Invasive Neurosurg*. 2004;47(1):16-23. doi:10.1055/s-2004-818347
29. Kassam AB, Prevedello DM, Carrau RL, et al. Endoscopic endonasal skull base surgery: analysis of complications in the authors' initial 800 patients: A review. *JNS*. 2011;114(6):1544-1568. doi:10.3171/2010.10.JNS09406