



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

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

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11

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

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

15

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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.

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## 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 <sup>1</sup>. Given that, several approaches to those regions have been developed through anterior, antero-lateral and postero-  
30 lateral routes <sup>2-4</sup>.

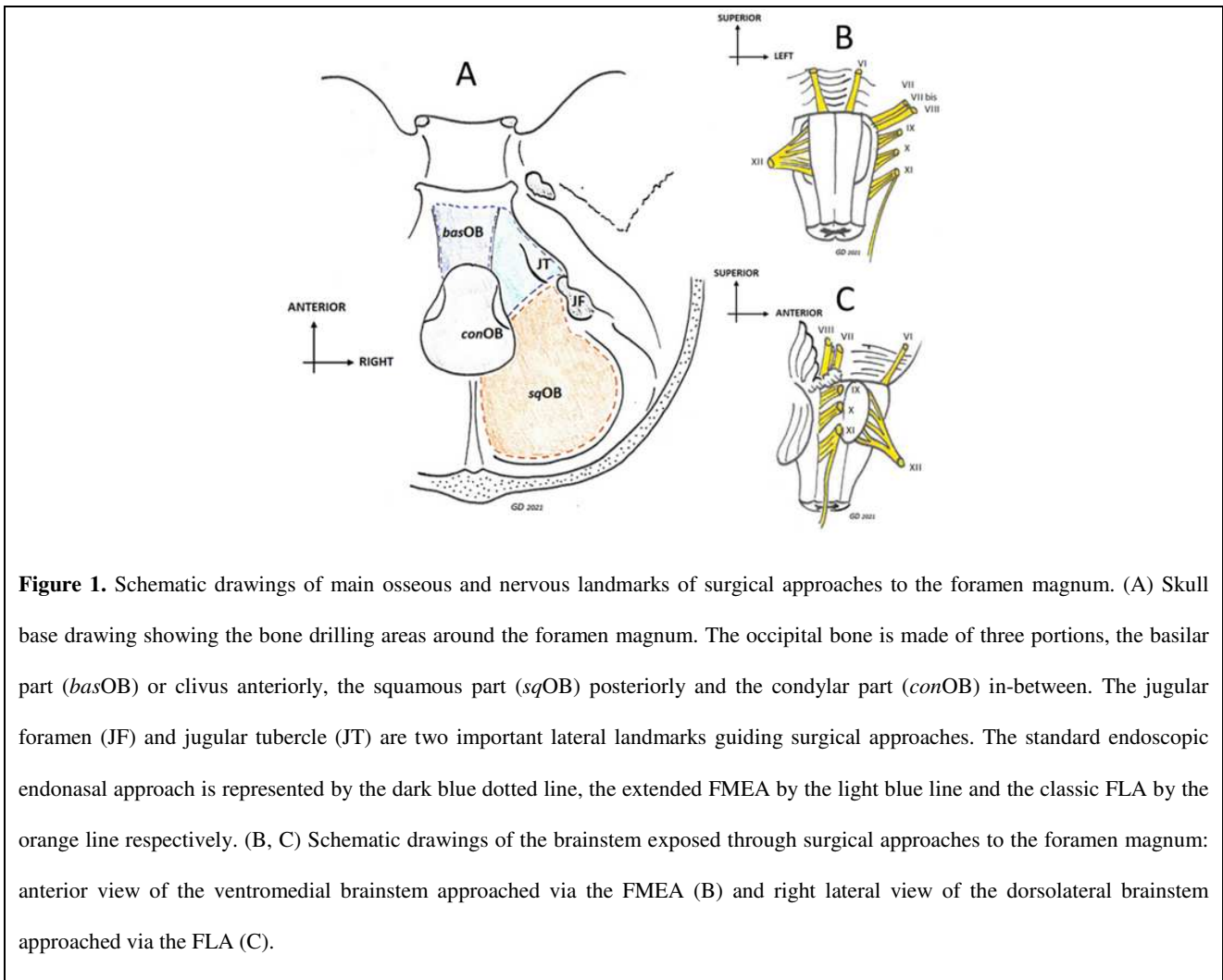
31 This variety of approaches reflects the difficulty to safely reach these regions. Neurosurgical corridors aim to provide  
32 the best possible exposure of the surgical field with the minimal manipulation or retraction of anatomical structures and  
33 therefore with limited morbidity. Nevertheless, 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 <sup>5</sup>, its popularity increased two years later with Bernard  
38 George series of 14 benign anterior FM tumours successfully resected <sup>6</sup>. 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 <sup>7-9</sup>, although some authors also have reported its use to manage intradural lesions <sup>10</sup>. The recent  
44 development of endoscopic endonasal approaches for the treatment of sellar region lesions <sup>11</sup> 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 <sup>12-16</sup>. 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 <sup>17,18</sup>. 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 <sup>19</sup>, 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.



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69 **MATERIAL AND METHODS**

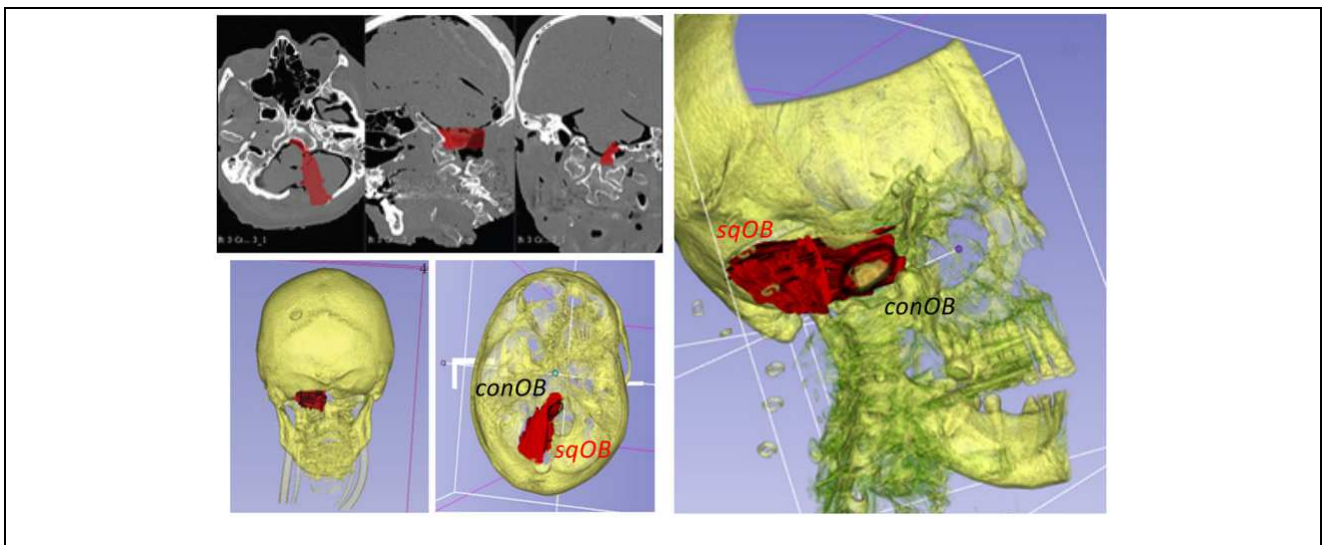
70 Dissections were performed on 10 fixed coloured silicone-injected adult head specimens, without previous brain or  
71 skull base abnormalities, at the Skull Base ENT-Neurosurgery Laboratory of the University Hospital of Strasbourg,  
72 France. Each specimen underwent a double FLA (for a total of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

74 Morphometric Analysis

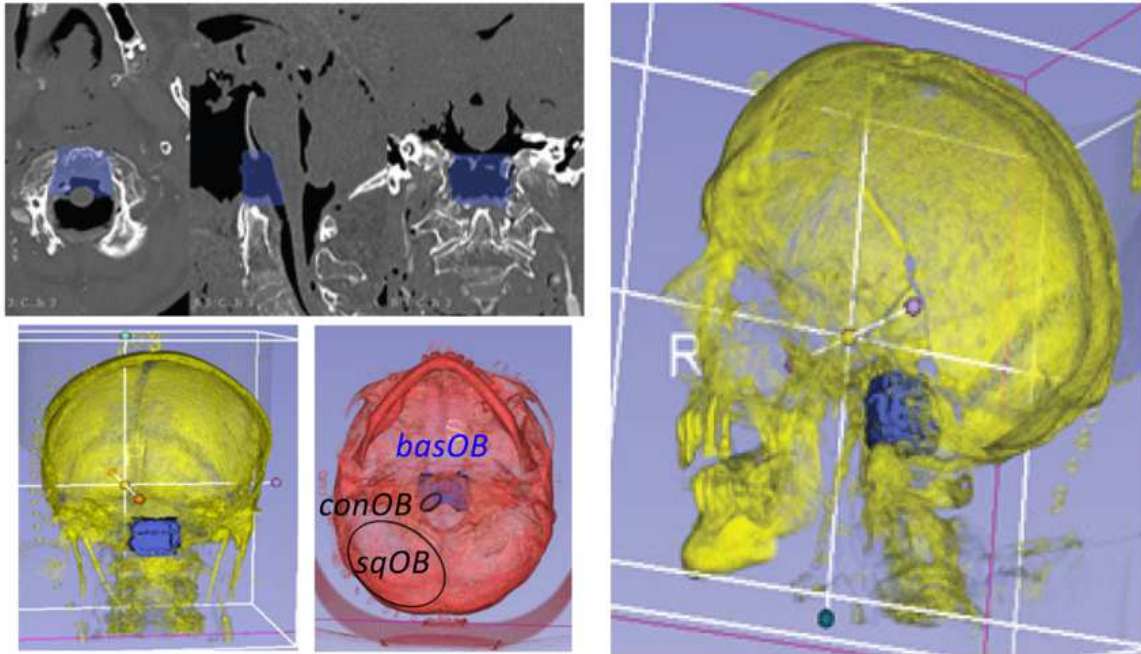
75 Before dissection, each specimen underwent a high-resolution computed tomography (CT) scan (1 mm cuts) to allow  
76 accurate measurements of deep anatomical structures exposed during dissection along the surgical corridor. Though  
77 MRI is the preferred imaging modality in clinical routine when 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.



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



92  
 93 **Fig. 3.** CT-scan with 3D reconstructions of the surgical volume involving the FMEA, accessing the foramen magnum  
 94 anteriorly through resection of the basilar portion of the occipital bone (*basOB*) highlighted in dark blue.

95  
 96 **Step-by-step description of surgical technique**

97 The FLA was performed bilaterally in all specimens, under microscopic magnification (Zeiss surgical microscope),  
 98 with the heads fixed in a three-pin Mayfield head holder, in a surgical position (i.e. prone position and slightly rotated  
 99 15° to the opposite side). Dissection was performed according to the techniques previously described in the literature,  
 100 consisting of three critical steps (Fig. 4) :

101 1°) A “hockey stick” skin incision is performed, extending from the spinous process of C3 to the inion, then curving  
 102 laterally toward the apex of the mastoid process. Muscles are dissected from the occipital bone and posterior arch of C1,  
 103 and retracted laterally. At the end of this first step, we identify the suboccipital triangle and the vertebral artery in its V3  
 104 segment.

105 2°) A C1 hemilaminectomy is done, extending from a point just beyond the midline to the groove of the vertebral  
 106 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.

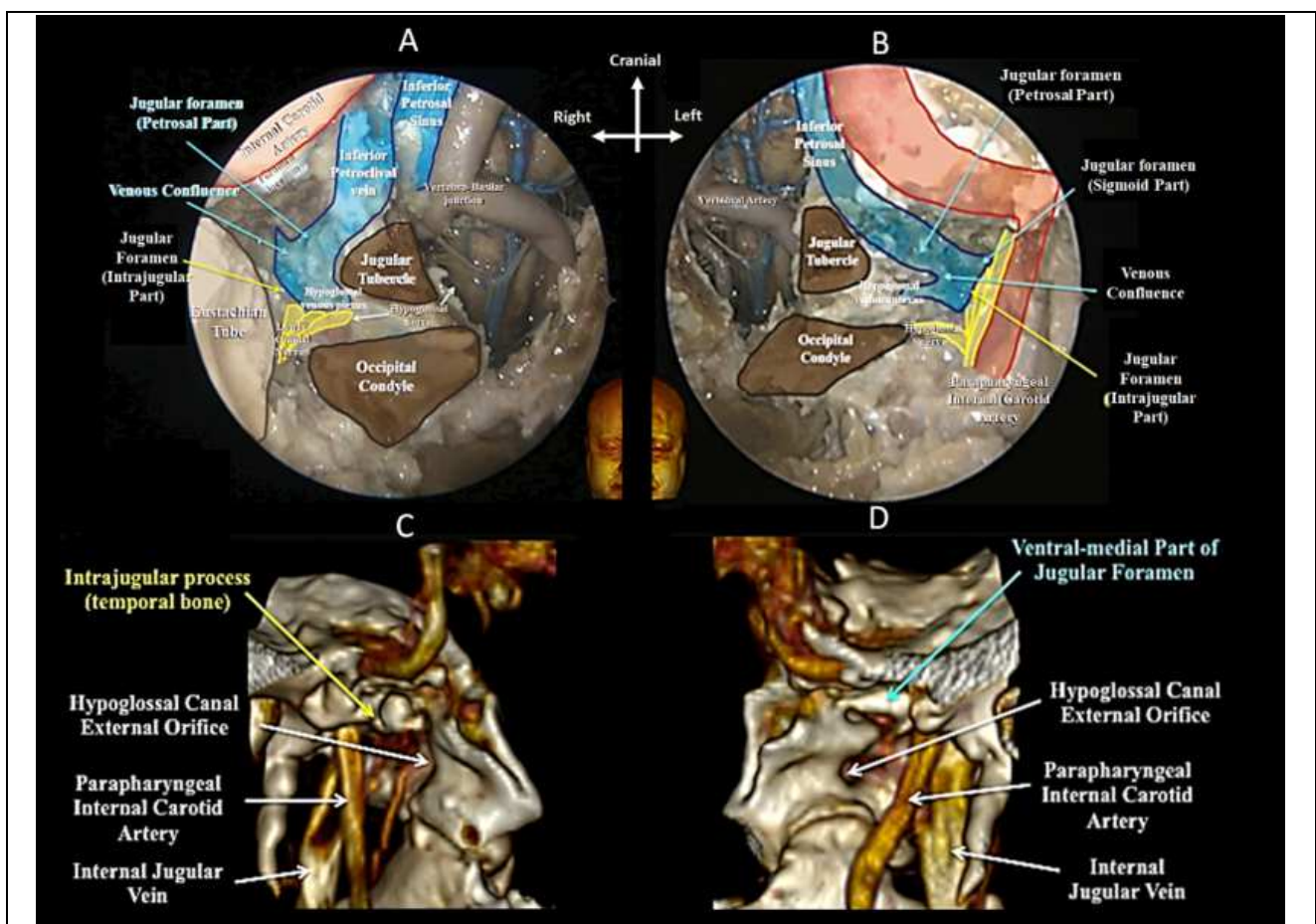
127 The FMEA is performed via a bilateral transnasal approach, through the following steps:

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.

144

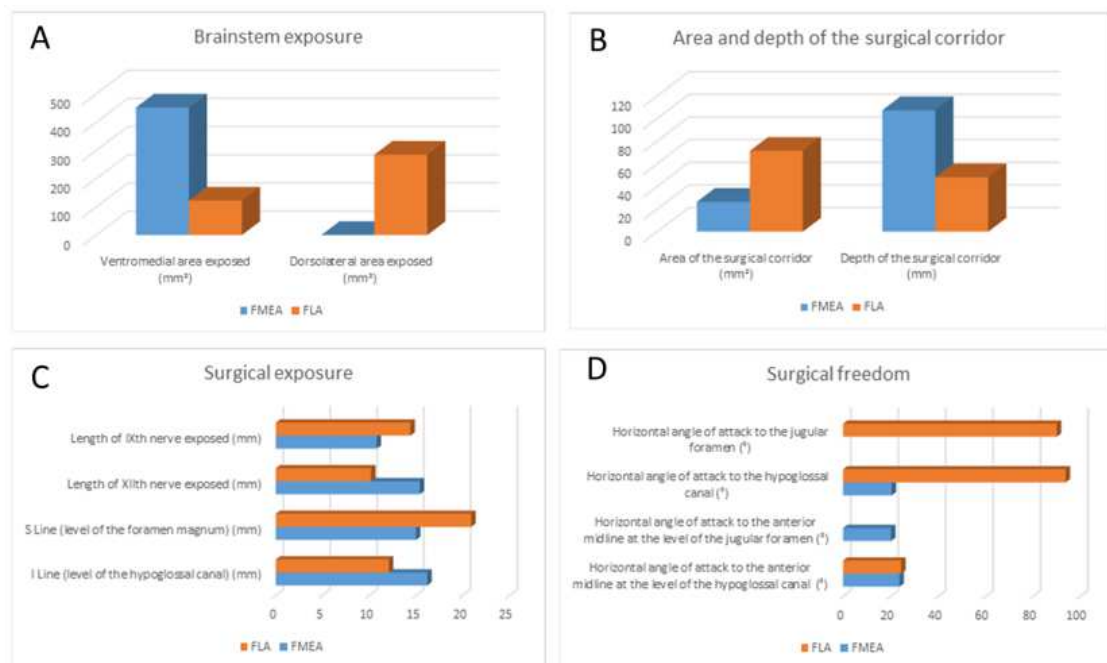
145 3°) Finally the dura mater is opened in the midline, allowing visualisation of the brainstem, the vertebral, basilar and  
146 origin of the posteroinferior cerebellar arteries, along with the cranial nerves IX, X, XI and XII bilaterally. (Fig. 5)

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  $180^\circ$  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  $107.2 \pm 3.70$   
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  $1.20 \text{ mm}$  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 ( $20.2^\circ$   
202  $\pm 1.48$ ). 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  $23.8^\circ$  and  $24.5^\circ$  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<sup>17,18</sup>.

### 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 corridors<sup>1</sup>. In the literature, several  
216 surgical approaches have been described for this region, each one with its pros and cons<sup>8,20</sup>. 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<sup>2,18,21</sup>.

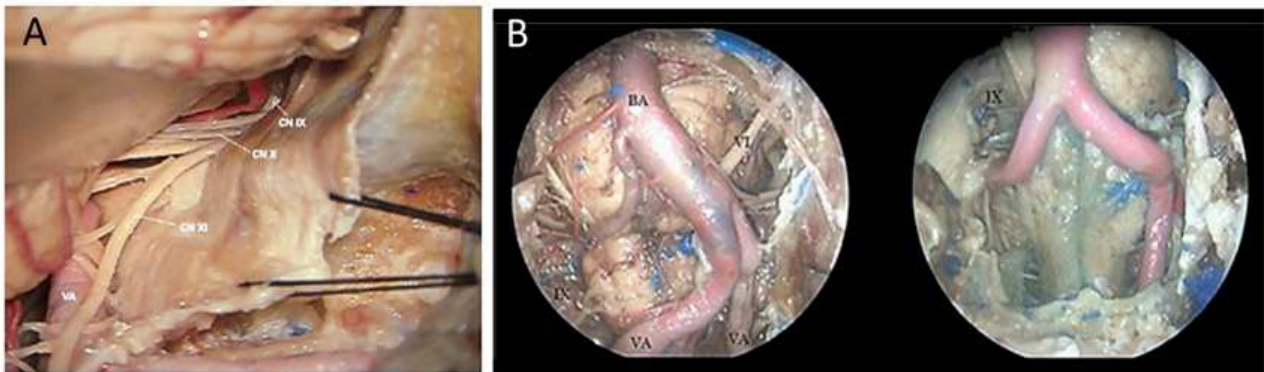
220 Having gathered as a group a 30-year experience with the FLA and a 15-year experience with the EEA <sup>4,16</sup>, we designed  
221 an anatomical study to compare the extent of surgical exposure and the freedom of movement (manoeuvrability)  
222 through both operating channels, FLA and FMEA.

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 <sup>3,20,22</sup>.

225 Most results from our anatomical study are in line with previous similar studies <sup>20,22</sup>, 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 <sup>9,23,24</sup>. 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 <sup>20,22,25</sup> (Fig. 7A). This approach also allows a wide exposure of the dorso-

243 lateral area of the posterior cranial fossa <sup>20</sup>. It should be pointed out, however, that the FLA offers a very limited access  
244 to the ventromedial brainstem area, in particular to the anterior midline at the cranial level. In fact, in our study it was  
245 possible to reach the clivus in only 30% of specimens due to the anatomical obstruction caused by a dominant VA;  
246 whereas at the level of the hypoglossal canal, the anterior midline could be reached in only 40% of specimens. Of note,  
247 at the level of the jugular foramen, the anterior midline could never be reached. Because of these limitations, the  
248 distance reached in a medial direction to the inferior clivus was of 21 mm at the level of the jugular foramen and of 12  
249 mm at the level of the hypoglossal canal.

## 250 **FMEA**

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

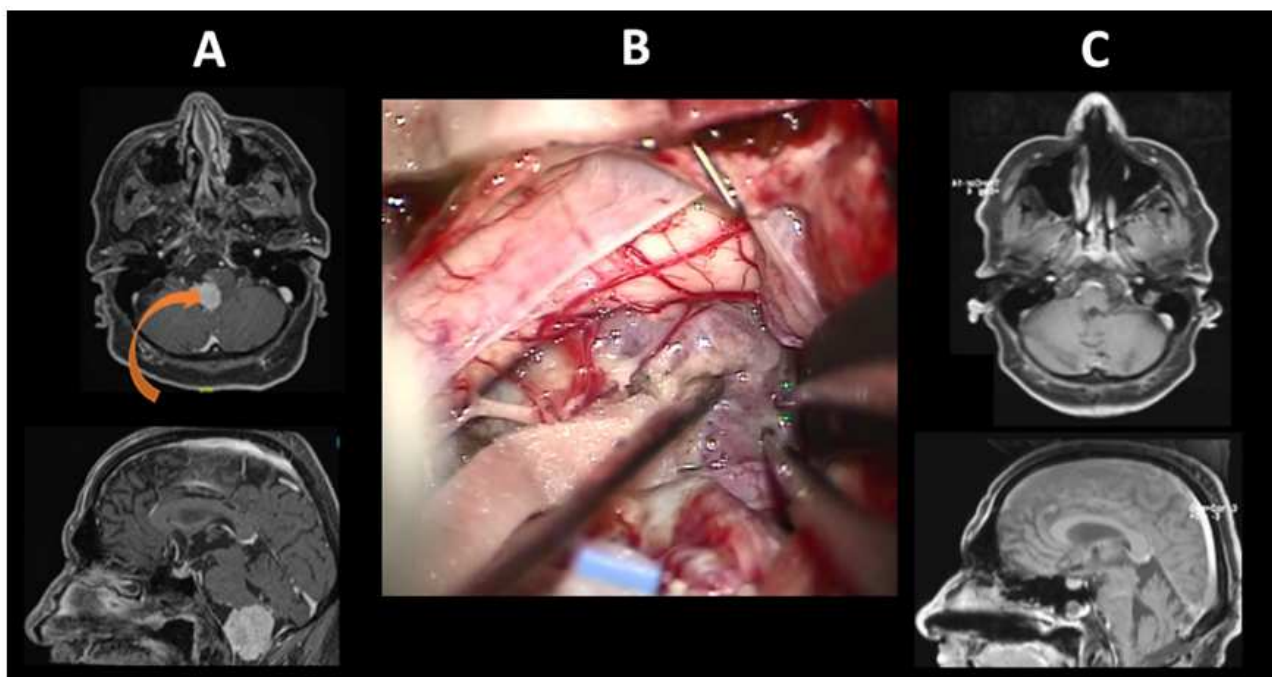
255 In our study, through FMEA the depth of the surgical corridor to the FM had an average of 10.5 cm and was  
256 significantly higher to that to the CCJ provided by FLA: because of the narrow surgical corridor, the FMEA is  
257 characterized by a lower grade of surgical freedom, compared to the open surgical approaches, in particular for what  
258 concern the laterality. Despite these limitations, our results showed that the FMEA allows an excellent exposure and a  
259 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  
260 midline of the jugular foramen, and of 1.8 cm from the hypoglossal canal. However, the lateral exposure of the lesion is  
261 limited by some uncrossable borders such as internal carotid arteries (ICAs), nerve of the pterygoid canal (vidian  
262 nerve), jugular veins, hypoglossal nerve and Eustachian tubes <sup>28</sup>. The degree of surgical freedom is also limited by the  
263 handling of the endoscope, because of anatomical obstacles such as the nose, the nasal concha and the depth of the  
264 endonasal corridor itself. Because of the above described limitations, the FMEA does not allow, in our opinion, a good  
265 exposure of lesions localized in the antero-lateral region of the FM and of the inferior clivus, in particular for those  
266 whose epicentre is localized at the level of the jugular foramen or at the hypoglossal canal. In fairness, the FMEA  
267 should not be considered as a mini-invasive approach because it is really destructive and can lead to important  
268 complications, such as the postoperative discomfort related to the turbinectomy, the risk of instability at the level of the  
269 CCJ and, most importantly, the risk of meningitis and CSF leak in case of intradural lesions <sup>29</sup>. According to our results,  
270 the FMEA is particularly useful for lesions with a small volume and near to the anterior midline (no further than 1.5 cm  
271 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 <sup>27</sup>. 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).



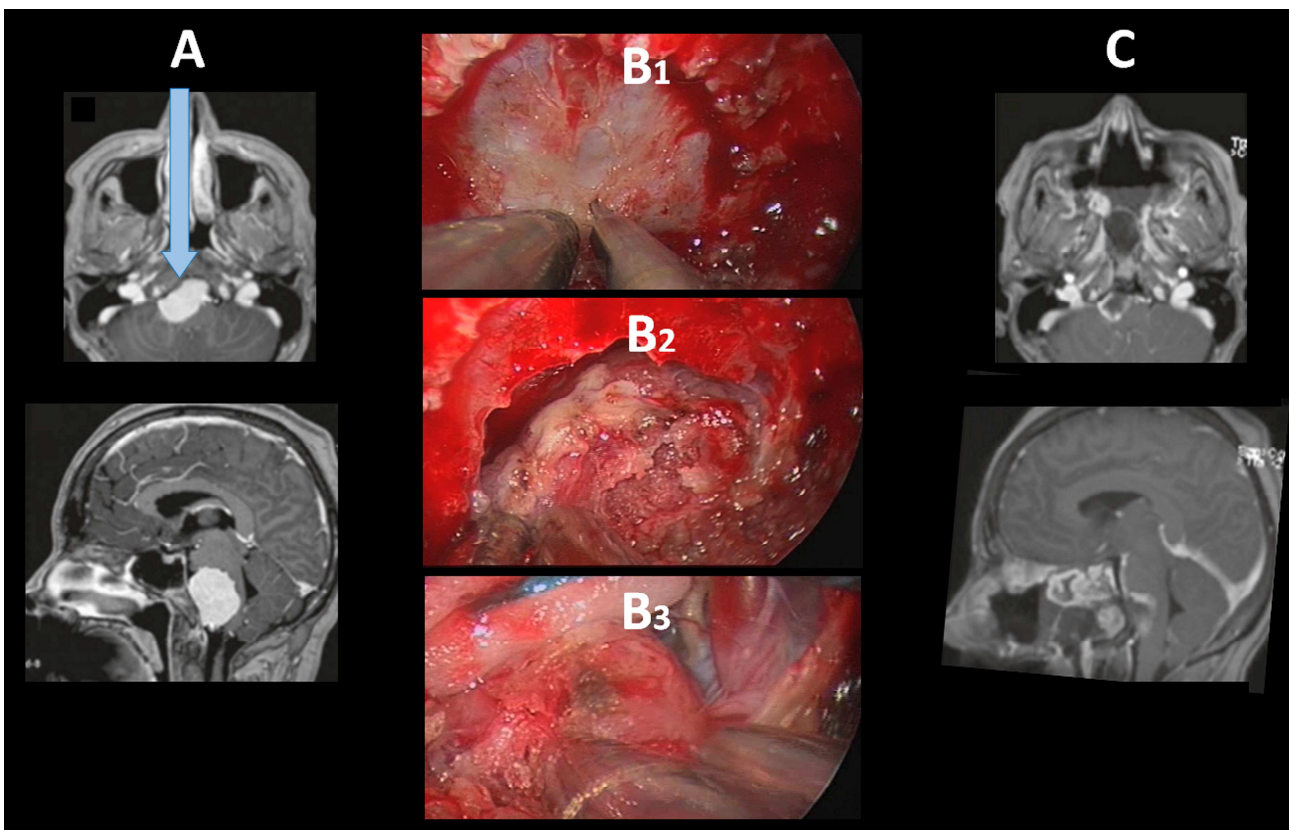
288  
289 **Fig. 8** Case vignette: FLA resection of a clival meningioma, preoperative MRI (A), perioperative view of FLA (B) and  
290 postoperative MRI showing complete removal (C).

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 (B<sub>1</sub>) followed by tumour exposure (B<sub>2</sub>) and visualization of leptomeninges and  
302 neurovascular structures along tumour resection (B<sub>3</sub>), 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.

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