Introduction

Published surveys show that unruptured aneurysms can be a real threat if left untreated\(^1\) and recent reports indicate a dramatic increase in the number of unruptured aneurysms being treated in the last years with favorable surgical or interventional outcome.\(^2,3\) When treating an incidental lesion with probable detrimental impact in the long-term, the outcome of our therapeutic strategies must outweigh the natural course of the disease. Hence, we should take every possible measure to reduce the morbidity caused by our therapy. In treating unruptured aneurysms, preservation of adjacent perforating arteries and maintaining blood flow in the parent vessel are of paramount importance. Various monitoring strategies are applied during an aneurysm surgery to perform a safe clipping such as recording motor evoked potential (MEP), intra-operative angiography, Doppler ultrasonography, and endoscopy.

Motor evoked potential monitoring during aneurysm surgery has been advocated and showed to predict postoperative neurological function.\(^4\) Changes in MEPs should be dealt with appropriately during temporary clipping of the parent artery or after permanent clipping of the aneurysm. Indocyanine

Role of endoscopy in multi-modality monitoring during aneurysm surgery: A single center experience with 175 consecutive unruptured aneurysms

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ABSTRACT

Objective and Background: Unruptured aneurysm surgery is a challenge to all vascular neurosurgeons as the patient is asymptomatic and no even slight neurological deficits should be expected postoperatively. To this aim, multi-modality checking of the vessels during the surgery is highly recommended to assure of the patency of the parent and perforator arteries next to an aneurysm. In this paper, we present our experience in the last 1.5 years with emphasis on the role of endoscope assisted microsurgery.

Methods: One hundred and seventy-five patients with unruptured intracranial aneurysms were operated in our institute in the last 1½ years. All patients underwent endoscope assisted microsurgery with pre- and post-clipping indocyanine green angiography. In selected cases, motor evoked potential monitoring was implemented.

Results: No mortality was observed in this period, and only 6 patients (3.4%) suffered new permanent neurological deficits postoperatively. Our illustrative cases show how endoscopy may help the surgeon to visualize hidden vessels behind and medial to an aneurysm.

Conclusions: Our results indicated that multi-modality monitoring during unruptured aneurysm surgeries is associated with excellent outcome. Endoscope is able to show blind corners during aneurysm surgery which cannot be routinely observed with microscope and its application in aneurysm surgery assists the surgeon to make certain of complete neck clipping and preservation of perforating arteries around the aneurysm.

Key words: Brain, clipping, endoscopy, monitoring, unruptured aneurysm

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green (ICG) video angiography (VA) is an alternative to the conventional intra-operative angiographies to evaluate the patency of the vessels and complete clipping. It has been previously reported how observing ICG under the microscope may change surgical planning to adjust an aneurysm clip. Sometimes, when the ICG-VA is repeated in a short time, the ICG has not washed out through the aneurysm and the clipped aneurysm also appears bright under the infra-red camera. We usually perform blood flow analysis with FLOW 800 software (Carl Zeiss, Oberkochen, Germany) in the clipped aneurysm to make sure there is no flow inside the sac after observing an almost constant intensity curve. Hence, FLOW 800 software is considered as a valuable addition to the monitoring armamentarium during aneurysm surgery. Endoscopic re-checking of the aneurysm neck and the clip is valuable in this scenario as well.

Unfortunately, ICG-VA visualization is limited to the vessels directly observed under microscope. To have a better view of the vessels hidden beyond the aneurysm, some authors have utilized Yasargil mirror in combination with ICG-VA. Another alternative to evaluate the hidden vessels is endoscopy. So far, endoscope-assisted microsurgery of aneurysms has become a routine practice in many centers, but only small case series have been published addressing its value in neurovascular operations. In one of these reports, 64 patients were included while 30 of them were ruptured and the rest unruptured aneurysms. To our best knowledge, the present study is the largest case series published so far which pertains to endoscope-assisted microsurgery of unruptured aneurysms.

Methods

A retrospective analysis of all patients with unruptured cerebral aneurysm operated in our hospital in the last 1½ years was performed. Patients’ demographic data, site of the aneurysm, operation notes, and postoperative morbidities or mortalities were reviewed and recorded. According to our department protocol, all patients diagnosed with unruptured cerebral aneurysms who are candidate for treatment, will be discussed with neuro-intervention team of our department. If not considered a good candidate for intervention, the patient will be offered surgery. We use reconstructed computed tomography angiography (CTA) routinely for surgical planning and in case of any ambiguity request a digital subtraction angiography. In our hospital, all neurovascular surgeries are performed with OPMI Pentero Microscope (Carl Zeiss, Oberkochen, Germany) with infra-red 800 camera equipped with FLOW 800 software (Carl Zeiss, Oberkochen, Germany). The camera allows us real-time intra-operative ICG-VA while the latter software provides hemodynamic information about blood flow. Upon exposure of aneurysm, we perform ICG-VA to clarify the aneurysm and its relation to all the surrounding vessels. Furthermore, to evaluate any perforating artery or other structures hidden behind the aneurysm we introduce a rigid endoscope (Machida, Japan) under microscopic guidance. With this technique, we check for the estimated final location of the aneurysm clip tips to be away from any critical structure. Technical nuances of aneurysm surgery have been discussed elsewhere in great detail. After applying the aneurysm clip (s), the vessels and the aneurysm are re-checked with ICG-VA. Whenever the distal blood flow is in question, blood flow measurements with Doppler ultrasound (DVM 4300, Hadeco, Japan) and FLOW 800 software are made. Once again, the endoscope is used to check for the final position of the blades and their relations to other structure and also to make certain that the aneurysm neck is completely obliterated. Based on the location of the aneurysm and probable risks, other monitoring systems such as MEP may be implemented. We do not use MEP monitoring for all patients although it has been advocated by some authors. Postoperatively, patients are kept for at least one night in neurosurgical intensive care unit under cardiac and neurological monitoring where any new neurological deficit is evaluated by CTA and/or magnetic resonance imaging (MRI) according to which necessary interventions are performed. Furthermore, CTA and CT-perfusion studies will be performed on 7th day postoperatively as a routine to evaluate any subclinical vasospasm and make sure of the proper placement of the clip.

Results

From January 2012 to June 2013, 175 patients with single unruptured aneurysm were operated by the senior author in our hospital. The mean age at diagnosis was 62.0 years (range of 24–79 years). There were 115 female and 60 male patients (female: Male ratio 1.92:1). Distribution of the aneurysms based on their site is shown in Table 1. The most common location of aneurysms was middle cerebral artery (MCA) followed by internal cerebral artery (ICA) and anterior cerebral artery (ACA). We observed no postoperative

<table>
<thead>
<tr>
<th>Site of aneurysm</th>
<th>Number</th>
<th>Percentage</th>
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<tbody>
<tr>
<td>ICA</td>
<td>58</td>
<td>33.2</td>
</tr>
<tr>
<td>IC-Oph</td>
<td>22</td>
<td>12.6</td>
</tr>
<tr>
<td>IC-PC</td>
<td>26</td>
<td>14.9</td>
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<tr>
<td>IC-Ach</td>
<td>6</td>
<td>3.4</td>
</tr>
<tr>
<td>IC-tip</td>
<td>4</td>
<td>2.3</td>
</tr>
<tr>
<td>ACA</td>
<td>44</td>
<td>25.1</td>
</tr>
<tr>
<td>AcCoA</td>
<td>36</td>
<td>20.6</td>
</tr>
<tr>
<td>DACA</td>
<td>8</td>
<td>4.5</td>
</tr>
<tr>
<td>MCA</td>
<td>69</td>
<td>39.4</td>
</tr>
<tr>
<td>Others</td>
<td>4</td>
<td>2.3</td>
</tr>
<tr>
<td>Total</td>
<td>175</td>
<td>100</td>
</tr>
</tbody>
</table>

ICA = Internal carotid artery; IC-Oph = Internal carotid ophthalmic segment; IC-Ach = Internal carotid choroidal segment; IC-tip = Internal carotid tip; ACA = Anterior cerebral artery; AcCoA = Anterior communicating artery; BA-Tip = Basilar artery tip; DACA = Distal anterior cerebral artery; MCA = Middle cerebral artery; IC-PC = Internal carotid communicating segment.
mortalities, but 6 patients suffered permanent and another ten transient morbidities during this period [Table 2]. Data for the patients who suffered permanent deficits are depicted in Table 3. All transient morbidities in 2013 were postoperative seizures fully controlled with anti-epileptic drugs. Postoperative CTAs did not reveal any neck remnant (unless deliberately) and no patients required further interventional or surgical treatment.

Discussion

Surgical outcome

Our results are in concordance with the recently published case series reporting 0.0–1.6% mortality and 6.2–9.4% morbidity for unruptured aneurysm clipping.\textsuperscript{[13–15]} In their recent review, Bacigaluppi et al. reported that a comprehensive intraoperative monitoring during aneurysm surgery reduces ischemia-related morbidities.\textsuperscript{[16]} We also believe that multi-modality intra-operative monitoring of the vascular patency before and after clipping is crucial for a safe aneurysm surgery. This is especially true of unruptured aneurysms where only a narrow margin of error is acceptable. Though endoscope is used commonly in many centers to check for a safe aneurysm clipping, our case series as the largest one addressing this issue, manifested its benefits in unruptured aneurysm surgeries with good outcome.

Advantages of endoscope assisted microsurgery

The main goals of a vascular surgeon while performing aneurysm surgery are to achieve a complete obliteration of the aneurysm while preserving normal neural and vascular elements. Endoscopy assists the surgeon to achieve both these goals [Figure 1] and its function turns vital in certain aneurysm locations when the aneurysm neck is usually surrounded by a bunch of perforators. These locations according to our experience and the literature include but not limited to the ICA-choroidal segment, ICA-communicating segment, MCA-M1, and MCA bifurcation. In one recent publication, the percentage of the different aneurysms with perforators originating from the neck was as follows: Basilar 7%, ICA bifurcation 17%, MCA main stem 12%, and anterior communicating artery (AComA) 11%.\textsuperscript{[17]} Although the author concluded that perforators originate infrequently from the aneurysm neck, these proportions are still so high that any negligence during the operation may increase morbidity of the patients dramatically.

M1 aneurysms have specific anatomical proximities which increase the chance of injury during surgical or interventional treatments.\textsuperscript{[18]} Aneurysms projecting superiorly are associated with lateral lenticulostriate artery (LSA) and frontal cortical branches while those projecting inferiorly accompany temporal cortical branches. Yeon et al. also showed that the closer an MCA-M1 aneurysm to the ICA bifurcation, the higher the chance of perforators injury.\textsuperscript{[19]} In another report by Iwama et al., the authors underlined the importance of the LSAs while operating a superior wall type aneurysm at the MCA-M1.\textsuperscript{[20]} Actually, we recommend and perform intraoperative MEP monitoring for all cases with this aneurysm location. Aneurysms in proximal MCA require special attention during localization and dissection of the aneurysm as some tiny perforators located at the posterior MCA wall and obscured by the temporal lobe may be damaged. In this scenario, endoscopy may come to help the surgeon with checking the final results after clipping.

Another location for the hidden perforators is the ventromedial wall of ICA when addressing posterior communicating artery (PComA) aneurysms particularly when the neck of the aneurysm extends medially [Figure 1a]. Wilson et al. reported their experience with the use of a Yarargil mirror during ICG-VA to elucidate medially located vessels in two cases with PComA aneurysms.\textsuperscript{[21]} PComA and anterior choroidal artery (AChA) are sometimes elusive but must be clearly visualized before and after clipping of these aneurysms as their injury may lead to the grave situation.\textsuperscript{[21,22]} Besides these two main branches by which a PComA aneurysm is flanked, there are important medially located perforators from PComA and ICA, which cannot be seen under microscope without significant retraction of the ICA and the aneurysm.\textsuperscript{[23,24]} By introducing an endoscope into the optic-carotid or carotid-oculomotor windows, one may visualize all these perforators and their relations to the aneurysm neck before and after aneurysm clipping [Figure 1d and e].

Surgery of AChA and ICA bifurcation aneurysms is also associated with an increased chance of morbidity, which
may be explained by their proximities to the early ACA-A1 and MCA-M1 perforating arteries. These locations are among the hot spots where endoscopy may assist the surgeon to identify perforators attached to the medial wall of the aneurysms and prevent morbidity.

Although the most frequent site of the aneurysm in our case series was MCA, none of these patients developed postoperative deficits. In fact, supracerebral ICA aneurysms composed the majority of neurological deficits. This finding was in accordance with the previous literature reviewed above.

Current literature

The first report of endoscope guided clipping in aneurysm surgery was published by Fischer and Mustafa. They used this method in 24 patients with 30 aneurysms and emphasized the benefits of endoscopy in improving the visualization of the pertinent anatomy. Taniguchi et al. reported benefits of endoscopy to clarify the anatomy in 81.5% of 44 cases operated for cerebral aneurysms. Interestingly, in 16.7% of their cases, the anatomy could be recognized only after endoscopy leading to clip adjustment in 9.3% of the surgeries. Wang et al. performed endoscopy-assisted microsurgery to treat 24 cases of ruptured aneurysms and found it very practical for providing the surgeon with good definition of surrounding structures such as cranial nerves, parent arteries, and perforators. In another report by Profeta et al. 52 patients with 58 aneurysms were operated with applying endoscope-assisted microsurgery over a period of 3 years. In four cases, endoscopy disclosed inappropriate clip placement requiring re-positioning or addition of an extra clip. The authors recommended endoscopy for certain aneurysm surgeries especially for those developed from the AComA.

Above-mentioned results should be interpreted cautiously. Usually studies evaluating a safety measure by performing an operation with and without a new tool (e.g., intra-operative MRI) in one session are biased in favor of the newly introduced device because the surgeon attempts to keep on the safe side by not exploring or removing the lesion without the safety device as he would do previously. This may improve the overall functional outcome but decreases intra-operatively measured goals before using the device (e.g., extent of tumor resection before performing intra-operative MRI). The ideal study for impact analysis is a prospective randomized controlled study, but as the technique we use here (i.e., endoscopy) is widely available, cost-effective, and with minimum harm to the patient it may be more ethical to rely on the results of such before-after studies.

Limitations and complications

Like any instrument, endoscopy has its limitations. Just confirming the correct placement of the blades is not always a proof of complete obliteration of the aneurysm. Conversely, sometimes even a partial clipping excludes the aneurysm from the circulation. Recent utilization of endoscopic ICG-VA may compensate for these shortcomings to some extent because it shows not only the external anatomy of the vessels but also the blood flow inside them.

When doing endoscope assisted microsurgery, usually the chief surgeon has to perform the endoscopy under microscopic guidance to avoid any unwanted damage to the surrounding structures, and the assistant surgeon will look at the endoscopy monitor. To come up with a solution, some companies offered a picture-in-picture mode for their microscopes. Furthermore, Hiramatsu et al. introduced a new endoscopic image display attached to the microscope eyepieces that allows the chief surgeon to watch both endoscopic and microscopic views almost simultaneously. We would like to videotape the endoscopic images rather than use either of the mentioned methods. The endoscopy is monitored by the assistant, and on its completion, the chief surgeon reviews the video if needed.

In spite of normal monitoring after clipping, sometimes a postoperative neurological deficit happens. Distal emboli during the vascular manipulation, irreversible brain ischemia due to prolonged proximal closure or permanent damage of perforators due to a misplaced clip can be the underlying causes. However, we still do not know whether discovering these events (i.e., emboli or irreversible ischemia) intra-operatively improves the outcome. To overcome these obstacles, we use intraoperative MEP especially when there is a high risk of distal emboli (e.g., severe atherosclerosis of the parent artery) or the aneurysm is placed in proximity to numerous perforators (e.g., proximal MCA aneurysms). This is in concordance with the studies reporting more difficult clipping and worse outcome for atherosclerotic calcified aneurysms. Vasospasm of a major or perforating artery can be another source of postoperative deficits despite a safe and uneventful clipping of the aneurysm [case 5 in Table 3] [Figure 2]. Although rare, but isolated symptomatic perforators vasospasm has been reported and it further signifies the importance of these vessels. Current technology does not offer surgeons an opportunity to monitor and prevent arterial spastic events during the operation, and in fact, the vasospasm related infarction develops in the postoperative period in most cases. In spite of aggressive medical or interventional management, the functional outcome is usually poor in this situation.

Another pitfall with both ICG-VA and endoscopy is that observing a patent vessel does not necessarily equal adequate blood perfusion. When suspected, we prefer to use FLOW 800 or Doppler ultrasound to measure distal blood flow. Though some efforts have been made to measure tissue perfusion with FLOW 800, its accuracy and reliability have not been proved so far.
Endoscope-assisted microsurgery may result in some complications as well, such as cerebral contusion or optic nerve damage.[7] The problem with endoscopy is that the surgeon may not view the structures superficial to the tip of the endoscope which may lead to inadvertent damage to the structures already passed by. This fact emphasizes the importance of a watchful monitoring of the endoscope lens under microscopic view.

**Purely endoscopic approach to aneurysms**

Although some authors have applied endoscopy for transcranial aneurysm dissection in addition to checking the safety of clipping,[36,37] we highly recommend microscope for dissecting the aneurysm: It gives us a three-dimensional view and high magnification which are indispensable advantages in aneurysm surgery. Furthermore, due to the probable complications discussed above, applying a purely endoscopic approach without microscopic monitoring might be perilous.

On the other hand, there are some reports of endoscopic endonasal approach to aneurysms.[38] We do not consider such approaches to be safe for several reasons. Intra-operative premature rupturing is always a concern even at the hands of experienced neurovascular surgeons.[8,15] Such a mishap may lead to a catastrophe in endonasal corridor as the intense bleeding obscures the view and securing a proximal control is very difficult if not impossible. Furthermore, current endoscopes do not provide surgeons with enough magnification to do microdissection around the aneurysm whilst it is an absolute necessity to dissect the perforators off the neck. Finally, though some progresses have been made in devising long instruments to perform trans-nasal surgeries, they are still less than perfect for microsurgical dissections.

We applied different monitoring tools to make a safe and secure aneurysm clipping as much as possible. Endoscopy demonstrates blind corners of the surgical field when different lenses are applied. Compared with ICG-VA, endoscopy can demonstrate the aspects that are concealed from our view normally. It is useful to show the position of clip and length of its blades, location of perforators before and after clipping, and the absence of the aneurysm neck remnant after clipping.

As discussed above, each monitoring device has its advantages and shortcomings during aneurysm surgery and only a multi-modality approach help the surgeon reduce the chance of possible morbidities and have a favorable surgical outcome.

**References**


