Indications for Surgery and Prognosis in Patients with Cerebral Cavernous Angiomas

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Abstract

Seventy-three cerebral cavernous angiomas were removed microsurgically from a series of 71 patients between August, 1983 and December, 1989. This retrospective investigation assessed the current indications for surgery and determined the prognosis for patients with cerebral cavernous angioma. There were 38 males and 33 females with a mean age of 37 years. Analysis included clinical presentation and history, neuroradiological findings, indications for surgery, and postoperative course. After an average follow-up period of 15 months, 35 patients were symptom-free, 16 had improved preoperative complaints, six were unchanged, and eight had deteriorated. Microsurgical extirpation of the malformation is indicated in all symptomatic patients where neuroimaging demonstrates the presence of a readily accessible cerebral cavernoma. Surgery is recommended in cases with deep-seated lesions causing massive hemorrhage, repetitive minor bleeding, or significant long-standing and progressive neurological disabilities. Clinically silent cavernomas located in eloquent regions of the brain contraindicate surgery, but should be closely monitored. Patients presenting with convulsions or neurological deficits caused by easily accessible cavernomas of the hemispheres have the best prognosis and a negligible risk for surgical complications. Those with deep-seated lesions of eloquent regions of the brain that have bled or caused sustained neurological disorders face the highest risk for morbidity owing to the surgical intervention, requiring careful preoperative evaluation.

Key words: cavernoma, cavernous angioma, arteriovenous malformation, surgical indication, prognosis

Introduction

Cavernous angiomas are a well known clinical and pathological entity.\(^6,22,27,25\) Recently, the incidence of detected congenital vascular hamartomas during life, and the reports on successful surgical treatment of cerebral cavernous angiomas have continuously increased.\(^1,2,5,7,8,11,14,21,26,29,30,33,34,36,39-42\) One of the main reasons is the high degree of sensitivity and specificity of high-field magnetic resonance (MR) imaging\(^4,17,19,20\) which is now the method of choice for preoperative evaluation of these benign and in most cases surgically curable vascular malformations.

The present investigation included 71 patients who underwent surgery during a 6-year period at our institution. The objective of this retrospective study was to investigate: 1) the current appropriate indications for surgery in cerebral cavernous angioma; 2) which patients definitely benefit from the surgical removal of a cavernous malformation; 3) which patients harboring a cerebral cavernous angioma must be considered at high risk for postoperative morbidity; 4) if surgery is clinically indicated, the optimal timing for the procedure in patients not presenting with massive hemorrhage. Special emphasis is placed on these questions because neurosurgeons are now dealing more frequently with cavernomas than in the past; and the incidence of hemorrhage and overall morbidity in the natural history of cavernous malformations are not entirely understood.\(^27,38\) Only recently have studies provided preliminary information about the risk of hemorrhage in patients with cavernous angioma.\(^4,19,20\)
Clinical Materials and Methods

I. Patient population

Seventy-one consecutive patients underwent microsurgical removal of symptomatic and histologically confirmed cerebral cavernous angiomas between August, 1983 and December, 1989 at the Department of General Neurosurgery, University of Freiburg. Two individuals harbored two cavernomas each at different locations which were treated surgically at different times. There were 38 males (54%) and 33 females (46%), ranging in age from 5 to 63 years (mean, 37 yrs). Familial occurrence was seen in two patients who were siblings.

II. Clinical presentation

Epileptic seizures were the only preoperative complaints in 29 patients. Another 16 combined seizures with various neurological deficits. Among the patients presenting with seizures, 21 suffered from generalized convulsions, seven from each of focal and complex partial seizures, and 10 from combined seizures. The incidence was low (one or up to three episodes of seizures in history) in 17 individuals, moderate (four seizures in history or several per year) in 13, and high (several seizures per month or more) in 15. Twenty-six patients received anticonvulsants, but the incidence of seizure was satisfactorily reduced in only 11 patients. The age, sex, and location, side, or size of the malformation did not significantly influence the incidence or the pattern of epileptic seizures.

Forty-nine patients presented with the clinical picture of a cerebral space-occupying lesion (Fig. 1). Progressive focal neurological deficits were manifest preoperatively in 44 individuals. The neurological disorders included impairment of cranial nerve function in 25 patients, sensory disturbances (chiefly hemihypesthesia and paresthesia) in 19, and motor deficits (predominantly hemiparesis) in 15. Severe headache, disorientation, dysphasia, dysarthria, trigeminal neuralgia, nystagmus, ataxia, and vertigo were additional signs and symptoms frequently encountered in these patients, depending upon the location of the malformation. Two patients with supratentorial cavernomas and three with lesions located in the quadrigeminal plate of the midbrain had symptoms of increased intracranial pressure. In the latter cases, this was due to obstructive hydrocephalus (Fig. 2), each requiring placement of a ventriculoatrial shunt prior to the removal of the vascular malformation. Cavernomas 2 cm in diameter or greater caused neurological deficits significantly more frequently than those less than 2 cm in diameter.

Four individuals presented with intracerebral mass hemorrhage from the malformation (Fig. 3). In another 39, less severe bleeding from the cavernoma...
was indicated by an ictal episode, confirmed intraoperatively. Approximately half experienced recurrent episodes of symptom exacerbation during the preoperative course, most likely caused by repeated microhemorrhages.

III. History
The period between first onset of complaints and surgery varied widely, ranging between 2 weeks and 26 years (mean, 49 mos). Seizures were of long-standing duration in most patients (mean, 65 mos). With one exception, patients harboring a pontine cavernoma had the shortest symptom duration (mean, 2.5 mos). Symptoms became manifest at a mean age of 33 years. Forty-four patients suffered a sudden onset of complaints. In the others, the symptoms developed gradually and most commonly progressed.

IV. Neuroradiological findings
Table 1 lists the various findings from preoperative computed tomographic (CT) scans in each patient. There were seven midline lesions (located in the cerebellar vermis, pons, or quadrigeminal plate of the midbrain), and 34 involving the right and 32 the left hemisphere. Table 2 displays the distribution of malformations according to location. Obviously, 47 lesions (64%) were quite easily accessible, whereas 26 (36%) were located deeply, in functionally important regions of the brain. The greatest CT diameter of the lesions ranged from 0.5 to 4 cm (mean, 1.9 cm).

Preoperative MR imaging in 60 patients was partic-

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**Table 1 CT findings of 73 cerebral cavernous angiomas**

<table>
<thead>
<tr>
<th>CT finding</th>
<th>No. of cases</th>
</tr>
</thead>
<tbody>
<tr>
<td>Hyperdensity</td>
<td>49</td>
</tr>
<tr>
<td>Mixed hypo- and hyperdensity</td>
<td>18</td>
</tr>
<tr>
<td>Ring enhancement</td>
<td>9</td>
</tr>
<tr>
<td>Hypodensity</td>
<td>2</td>
</tr>
<tr>
<td>Calcifications within the lesion</td>
<td>29</td>
</tr>
<tr>
<td>Perilesional edema</td>
<td>10</td>
</tr>
</tbody>
</table>

**Table 2 Location of 73 cavernous angiomas**

<table>
<thead>
<tr>
<th>Location</th>
<th>No. of cases</th>
</tr>
</thead>
<tbody>
<tr>
<td>Supratentorial</td>
<td></td>
</tr>
<tr>
<td>frontal</td>
<td>21</td>
</tr>
<tr>
<td>temporal</td>
<td>14</td>
</tr>
<tr>
<td>parietal</td>
<td>8</td>
</tr>
<tr>
<td>occipital</td>
<td>3</td>
</tr>
<tr>
<td>insula</td>
<td>6</td>
</tr>
<tr>
<td>basal ganglia</td>
<td>4</td>
</tr>
<tr>
<td>thalamus</td>
<td>2</td>
</tr>
<tr>
<td>Infratentorial</td>
<td></td>
</tr>
<tr>
<td>pons</td>
<td>8</td>
</tr>
<tr>
<td>midbrain</td>
<td>5</td>
</tr>
<tr>
<td>brachium pontis</td>
<td>1</td>
</tr>
<tr>
<td>cerebellum</td>
<td>1</td>
</tr>
</tbody>
</table>

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Fig. 3 Postcontrast CT scans of a 30-year-old male. Preoperative scans show a ring-shaped lesion within the dorsal pons (left). During observation, he developed sudden onset of right hemihypesthesia, dysarthria, and abducens nerve paresis due to spontaneous bleeding from this pontine cavernoma (right). The lesion was completely extirpated through the fourth ventricle. At 3-year follow-up, he was symptom-free.
ularly useful in confirming the suspected diagnosis of cavernous angioma and in delineating the relationship of the malformation to the surrounding brain structures (Figs. 4 and 5).

Cerebral angiography was performed in each case. A normal angiogram was found in 50 cases. A subtle but definite vascular stain was evident in only two pontine cavernomas. Filling of prominent draining veins adjacent to the cavernoma was seen in four frontal, and one each of temporal, occipital, insular, and pontine malformations, suggesting a venous angioma.

V. Indications for surgery

The indications for surgical intervention in this series included treatment of convulsions, improve-
ment of focal neurological disorders, correction of cerebrospinal fluid pathway obstruction, evacuation of hematoma, and prevention of further bleeding. The latter was a prophylactic surgical indication. Eliminating the risk of a potentially life-threatening massive bleeding was appropriate mainly in patients with repetitive hemorrhages or with lesions of eloquent areas of the brain, where even minor bleeding had already caused substantial morbidity.

Superficially located malformations were reached by exposing the convexity of the brain. More deeply placed lesions were generally approached by dissecting the cerebral sulci or fissures, and removed from the closest point to the surface of the brain or ventricles. \(^1\) The surgical approaches used in this series are presented in Table 3.

### Results

#### I. Early postoperative course

Table 4 summarizes the overall surgical outcome by discharge (10-14 days after surgery in 61 cases). Of the patients not presenting with neurological disorders, only one experienced transient deficits following the procedure which had resolved by follow-up examination. The surgical complications are listed in Table 5. A detailed analysis of the complications encountered in patients with deep-seated malformations has been presented elsewhere. \(^1\)

#### II. Follow-up

Follow-up data were available from 65 individuals (92%), obtained from questionnaires, telephone contact, or re-examination. The mean follow-up period was 15 months, ranging from 8 months to 5 years.

The evolution of the seizure disorder by follow-up examination is presented in Table 6. Excluding the first 5 postoperative days, 27 patients presenting with epileptic seizures remained seizure-free, either with or without antiepileptic medication. The prevalence of seizures clearly decreased from 60% preoperatively to 18% postoperatively. Additionally, the incidence was markedly reduced in 75% of the 12 patients still suffering from convulsions. These 12 individuals had a significantly longer duration of symptoms (mean, 124 mos) compared with the seizure-free patients (mean, 34 mos). The size of the lesion did not influence the outcome.

Table 7 summarizes the evolution of the main neurological disorders. In 15 of the 22 patients with...
unsatisfactory neurological condition in the early postoperative stage, deficits had resolved or markedly improved by follow-up examination.

Overall, by follow-up examination, 35 individuals (54%) were symptom-free, 16 (25%) had improved preoperative complaints, six (9%) remained unchanged, and eight (12%) had deteriorated.

**Table 7 Evolution of main neurological deficits by follow-up**

<table>
<thead>
<tr>
<th>Neurological deficits</th>
<th>No. of patients*</th>
<th>Cured</th>
<th>Improved</th>
<th>Unchanged</th>
<th>Worsened</th>
<th>Unknown</th>
</tr>
</thead>
<tbody>
<tr>
<td>Motor</td>
<td>15</td>
<td>5</td>
<td>3</td>
<td>3</td>
<td>3</td>
<td>1</td>
</tr>
<tr>
<td>Sensory</td>
<td>19</td>
<td>7</td>
<td>5</td>
<td>3</td>
<td>2</td>
<td>2</td>
</tr>
<tr>
<td>Cranial nerves</td>
<td>25</td>
<td>10</td>
<td>5</td>
<td>6</td>
<td>3</td>
<td>1</td>
</tr>
</tbody>
</table>

*Preoperatively.

Discussion

Age and sex distribution, history, and clinical and neuroradiological findings in this series compare favorably with details given in other reports. The MR imaging features of cavernomas may not be pathognomonic, but a correct diagnosis was possible in virtually all our patients for whom preoperative MR images were obtained. As in other reports, MR imaging offered several advantages: a high degree of specificity; detection of even very small lesions, particularly within the posterior fossa; and sufficient accuracy of triplanar sections to determine the exact location of the malformation and the relationship with the surrounding cerebral structures.

Cerebral angiography is not primarily a diagnostic tool, but can eliminate coincidental vascular malformations, such as cavernous angioma associated with venous angioma or capillary teleangiectasia, both rare but possible occurrences. MR imaging offered several advantages: a high degree of specificity; detection of even very small lesions, particularly within the posterior fossa; and sufficient accuracy of triplanar sections to determine the exact location of the malformation and the relationship with the surrounding cerebral structures.

Assessment of appropriate indications for surgery upon cavernous angiomas located within functionally important regions of the brain, however, remains a constant challenge. Such lesions may remain clinically silent, and are now usually detected by chance on MR images. They may cause various symptoms due to repeated microhemorrhage or to gradual enlargement, or they may suddenly cause the patient's condition to become critical and life-threatening when massive hemorrhage occurs. Controversy continues over whether surgical or conservative treatment is preferable in such cases. Previously, surgical removal of deep-seated lesions, particularly of those located within the brainstem, has been carried out mainly in patients with local massive bleeding, or with recurrent subarachnoid hemorrhage. We also consider that extirpation of the malformation is the therapy of choice in such cases, in order to avoid a second, potentially fatal bleeding.

Apart from cases with major bleeding, we operated on patients with cavernomas in sensitive regions of the brain presenting with various symptoms other than massive hemorrhage from the malformation. Only a few similar cases have been published. In the initial period, our results were sometimes disappointing. Later, our surgical success rate has clearly increased, and serious complications have become a rarity since 1988. Encouraged by these results, and like several authors, we now prefer a rather aggressive surgical policy in patients presenting with either repetitive minor bleeding (and correlating symptoms) or with sustained hemorrhages. Although the follow-up period is not long enough to allow definitive inferences, the vast majority of our patients presenting with convulsions experienced a significant amelioration of their seizure disorder postoperatively. Similarly, those presenting with neurological disabilities caused by cavernous angiomas located in noneloquent regions of the brain unequivocally benefited from surgery. Surgical therapy was less effective, however, in patients suffering from uncontrollable long-standing seizures.

Authors have been unanimous for more than a decade in considering surgery the therapy of choice in readily accessible cavernous angiomas causing clinical symptoms and/or recurrent hemorrhages. We now prefer a rather aggressive surgical policy in patients presenting with either repetitive minor bleeding (and correlating symptoms) or with sustained hemorrhages. Although the follow-up period is not long enough to allow definitive inferences, the vast majority of our patients presenting with convulsions experienced a significant amelioration of their seizure disorder postoperatively. Similarly, those presenting with neurological disabilities caused by cavernous angiomas located in noneloquent regions of the brain unequivocally benefited from surgery. Surgical therapy was less effective, however, in patients suffering from uncontrollable long-standing seizures.

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neurological deficits, provided that the malformation extends close to or reaches the surface of the brainstem or ventricles, including the surface of the third ventricle. We must be aware, however, that these individuals face the highest risk for transient and even permanent morbidity attributable to the surgical intervention. A single episode of minor bleeding, or less severe symptoms which rapidly resolved, is not necessarily a surgical indication upon deep-seated lesions.

In contrast to arteriovenous malformations whose natural history is now well defined, the yearly risk for massive hemorrhage from cavernoma is difficult or impossible to precisely predict. Del Curling et al. recently estimated the annualized bleeding rate at 0.25%/person-year of exposure, and Robinson et al. 0.7%. The risk of rebleeding after an initial hemorrhage, however, remains unknown. Nevertheless, there is no doubt that these lesions can be the source of major hemorrhage, and the consequence of such an episode in a patient harboring a deep-seated cavernoma may be devastating. A careful evaluation of the optimal therapy is therefore mandatory in patients with a symptomatic deep-seated cavernoma not presenting with massive bleeding. There is no indication for surgery in patients with clinically silent cavernous angiomas located in critical regions of the brain, particularly when approaching these lesions would require traversing healthy and functionally important cerebral structures. Certainly, such patients need to be carefully followed clinically and neuroradiologically at least once per year.

Acknowledgments

Dr. Bertalanffy is the recipient of a fellowship from the Japanese Society for the Promotion of Science, Tokyo, Japan, and the Alexander-von-Humboldt Foundation, Bonn, Germany.

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