

## Case Report

# Meningeal Anaplastic Hemangiopericytoma

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## Case Presentation

The case features a 66 year old woman with a history of high blood pressure during pharmacological treatment. She had no other pathological history. She presented with a right hemicranial headache which had become resistant to regular analgesics. The physical exam, including neurological and eye fundus tests, showed no abnormalities.

The Computerized Tomography (CT) scan showed a right temporal lesion which was hyperdense, with hypodense centre and marked perilesional edema, compression of the right lateral and third ventricles, as well as subfalcine herniation with intense enhancement after administration of contrast. In the Magnetic Resonance Scan (MRI), we saw a lesion of 55x45x45mm, with deformity of the Sylvian fissure. This was an isointense T1 lesion, with the central area being of less intensity, markedly heterogenous in T2 Flair and spin echo sequences, leaving the hyper intense central areas – which describe a variegated morphology – visible. This coincided with the areas of greater magnetic susceptibility in T2 sequences, attributable to the presence of necrosis and vascular structures. In the periphery of the lesion, we could see a displacement of the vessels, with no signs of infiltration. This was surrounded by a digit form vasogenic edema, which causes displacement of the parenchymatous structures of the basal ganglions and of the middle line, with the ipsilateral ventricle being collapsed. After the intravenous administration of gadolinium, the lesion was intensely enhanced, leaving a non-captant central area, with no clear “dural tail”, although it appeared to be in contact with the lesion at its posterior part (Figure 1). The perfusion sequence reveal great lesion vascularization, with a CBV always greater than x4, at some points even reaching x12. Spectroscopy didn't offer any specific defining characteristics.

Prior to the surgical resection we carried out tumor embolization. We could see arterial irrigation in the tumor, both through the ICA and in the branches of the external CA on the right side (mainly via the ECA), with venous draining towards the sigmoid sinus.

The patient underwent a complete microsurgical in bloc resection of the extra-axial lesion, via the arachnoid plane. There were no complications during the procedure. The postoperative MRI scan carried out after surgery didn't show any tumor remains. The symptoms passed after surgery.

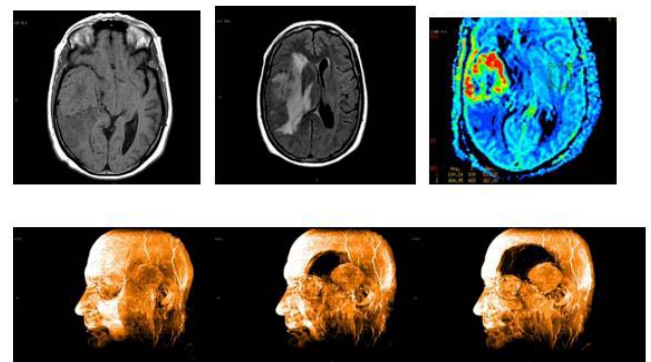
### Abstract

Hemangiopericytoma (HPC) is a rare tumor with high risk of recurrence at neural and metastatic potential level. We report the case of a woman with a primary Meningeal Anaplastic Hemangiopericytoma (MHPC) at right temporal lobe. In the same way, given the low frequency of this tumor, we have reviewed the available data on its characteristics, as well as its therapeutic management.

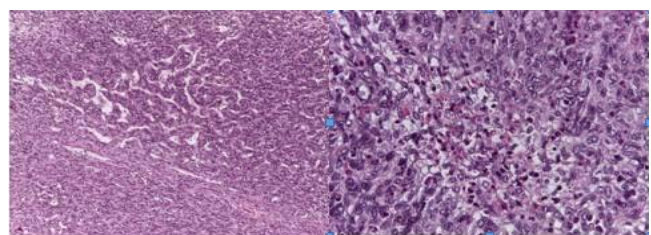
**Keywords:** Hemangiopericytoma; Intracranial meningeal neoplasm

The hystopathological examination revealed hyper cellular tumors composed of fusiform, pleomorphic cells, arranged in all directions. They featured focus of ischemic necrosis and 6x10 mitotic activity, fields of great increase. Irregular vessels could be seen, focally appearing to be of “staghorn” nature. Some vessels were of great size and still bore this appearance, with chemo-embolization material being noticed, together with the main ischemic foci. Tumor cellularity was surrounded by a web of abundant reticulin (Figure 2).

Immunohistochemical staining revealed positive staining for CD34 (focal), vimentin, reticulin and bcl-2, and negative for Epithelial



**Figure 1:** Pre-operative magnetic resonance images show an extra-axial mass in the temporal lobe (A) T1- weighted, isointense multilobulated margin, and deformity of the Sylvian fissure, (B) T2 –weighted, compressing of the right lateral and third ventricles with surrounding edema. (C): Perfusion sequence reveal great lesion vascularization (D). 3D magnetic resonance scan.



**Figure 2:** Hematoxylin and eosine stain reveals the pathology of the tumor is a diagnosis os meningeal hemangyoperitoma. hypercellular tumor composed of pleomorphic cells, Irregular vessels focally appearing to be of “staghorn” nature.



**Figure 3:** Immunohistochemistry. Tumor cells strongly expressing vimentin (A), focal CD 34 (B), and reticulin (C).

Membrane Antigen (EMA), Glial Fibrillary Acidic Protein (GFAP), CK AE1 / AE3 and S-100. The proliferation index, examined via the Ki67 antibody, reached 8-10% (Figure 3).

After the procedure, the patient underwent Volumetric Modulated Arc Therapy (VMAT technique) for three months on the surgical bed. This was finalized by May 2014, with the patient receiving a total doses of 58Gy. Twelve months after the surgical resection, no clinical-radiological evidence of local recurrence existed (Figure 4), nor did any evidence of distant metastases.

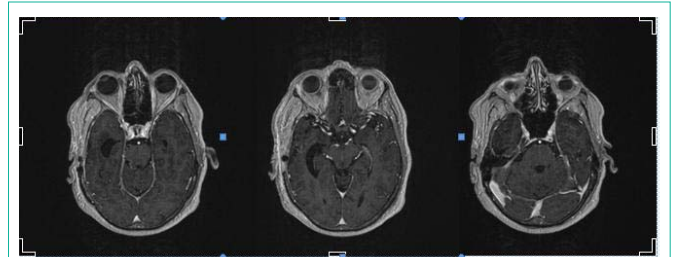
## Discussion

The Hemangiopericytoma are uncommon hypervascular neoplasms, of mesenchymal lineage. They arise from the malignant transformation of Zimmerman pericytes – contractile spindle cells surrounding capillaries and post capillary venules. HPCs may occur anywhere in the body; usually the most commonly reported locations are lower extremities and retroperitoneum. Intracranial HPC constitute 2-4% of meningeal tumors and less than 1% of the tumors in the central nervous system [1-5].

The hemangiopericytoma was first reported by Stout and Murray in 1942 [6], and the first reported case of hemangiopericytoma originating in the meninges was described by Begg and Garret in 1954 [7]. They were originally considered a variation of meningioma, and they were given the name of angioblastic meningioma. More recently, thanks to its special clinical behavior, together with its immunohistochemical, structural and genetic characteristics, it has been recognised as a different entity altogether [8-12]. Since 1993, the World Health Organization's (WHO) [13] classification of tumors in the nervous system recognizes them as being nonmeningothelial mesenchymal tumors.

The interest in differentiating meningiomas from hemangiopericytomas – with which they share similar clinical and radiological characteristics – resides in the fact that hemangiopericytomas are characterized by their propensity to local recurrence and the potential to metastasize in extraneural areas [2,4,14]. Thus, given their aggressiveness, they need a selection of optimal treatment.

The chosen treatment for intracranial hemangiopericytomas is of a surgical nature. A total resection of the lesion is recommended whenever possible, due to the fact that the resection has correlated with the free interval of recurrence and overall survival [1,3-5,14,15]. Given that they are very vascularized tumors, with predisposition to bleed heavily during operation, pre-surgical embolization must be considered [3,4]. Nevertheless, a complete surgical resection does not eliminate the high risk of recurrence and in the majority of cases, recurrences that occur in the central nervous system do so in the place



**Figure 4:** Post-operative magnetic resonance images taken within 12 month of surgery show no evidence of local recurrence.

where the original tumor appeared [2,16]. The most important factor which determines the recurrence and prognosis in these patients is the extension of the surgical resection [1,2,15]. There are studies which show that a total resection of the lesion prolongs the moment of recurrence for an average of 65 months [2,17]. Furthermore, a meta-analysis has demonstrated a survival ratio of 10 years through 69% of those who underwent a gross total resection, compared to 44% of those who obtained a subtotal resection [18].

Treatment with adjuvant radiotherapy is controversial, since the data we have available originate in small samples of retrospective studies which have conflicting results. Studies do exist with postoperative radiotherapy, radiotherapy with doses of at least 45-50 Gy (since the HPC sensitivity is dose-dependent) [19,20] even in cases of complete resection, which has proven useful in increasing recurrence-free interval [12,21], although this therapy has not shown to bear effect on recurrences along the neuraxis or against distant metastases [12,21]. On the other hand, there are studies which show no benefit in the addition of radiotherapy after a total resection [18]. However, it seems logical to suggest the use of adjuvant radiotherapy and cases of incomplete resection – either in lesions that are technically unresectable, or as an exclusive palliative treatment. Radio surgery seems to be an effective treatment with favorable results for recurrent or residual MHPC that are well-defined and small in size [1,2]. As well as having a tendency for local recurrence and recurrence along the neuraxis, intracranial hemangiopericytomas also tend to metastasize in other areas outside of the nervous system, and they are able to appear several years after the treatment of the primary tumor [22,23], which makes it necessary to have follow up monitoring lasting as long as possible. Bone, liver and lung are the most commonly reported sites of metastasis in HPC. Additional resections must be considered in these circumstances, although this is not always possible [24]. A strategy of optimal treatment of advanced HPC has not yet been identified [14,24-26].

The role of chemotherapy based on anthracyclines is still controversial, with modest efficacy in the treatment of MHPC [27,28]. Recent studies have identified the fusion gene NAB2-STAT6, both in HPC and in solitary fibrous tumors [29,30]. Although this hasn't yet been translated into therapeutic targets, it certainly does suggest a new line of research. Hypervascularization of these tumors, and their immunohistochemistry (IHC) expression of the Vascular Endothelial Growth Factor Receptor (VEGFR) and Platelet-Derived Growth Factor Receptor (PDGFR) [19,31-33,34,35] is the rational basis for encouraging the study of targeted therapies with anti-angiogenic drugs. Until now, only small experimental studies and a series of reported cases have been published. Some of these propose

the possible role of IFN- $\alpha$  [22,36] in the stabilization of the disease. Others suggest that imatinib [34,35], sorafenib [34] and sunitinib [37,38] can achieve stable, sustained responses. There have also been cases reported with pazopanib [39], a multi-targeted tyrosine-kinase inhibitor [40]. This showed efficacy in patients who had a metastatic hemangiopericytoma, with longer-term recurrence free survival and overall survival [41,42]. There is a retrospective study in which the combination of temozolomide and bevacizumab has a high rate of partial Choi responses, making it a promising therapy [28]. Despite these results, further research is needed for validation in other prospective studies.

## Conclusion

Given the high risk of recurrence, the MHPC must be completely resected, as long as it is technically possible. Even still, when a complete resection is performed, adjuvant RT should be considered, as this appears to reduce the rate of local recurrence. Long-term monitoring is essential to detect recurrences or distant metastases, which can appear decades after the correct handling of an initial primary tumor.

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