Stereotactic Radio Surgery for Residual and Recurrent Central Neurocytoma

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Abstract

Background: Central neurocytoma (CN) is a rare benign tumor of central nervous system that is typically located in the lateral ventricle and constitute 0.1–0.5% of all primary brain tumours.

Methods: We are relating retrospectively the results of 11 patients who underwent Gamma knife radio surgery for CNs in the Gamma unit at functional and stereotactic neurosurgery service in Marseille from January 1999 to December 2015. The mean age at Gamma knife radio surgery (GKS) was 25.12 years (range 17-41 years). All patients underwent surgery before (GKS) with a mean delay of 19, 37 months (6-62 months). The prescribed mean dose delivered to the tumour margin was 19, 1 Gy (12-24 Gy). The mean volume of treatment was 4.4 mL (1.03-10.7 mL). The mean time follow-up was 55.2 months (7.2-108 months).

Results: The local tumor control was 90% at the final follow-up time. One recurring tumour was local (4.67 years after GKS) and 2 were “out-of-field” recurrence (5.42 and 6.8 years after GKS). Repeat stereotactic radio surgery was performed in two patients with an average delay between the two of 5.91 years and an average dose of 14.34 Gy. The local tumour control was also unrelated to the prescribed dose (p=0.89) and the tumour volume (p=0.67). No deaths and no radiation injury were observed during our follow-up.

Conclusion: Gamma knife radio surgery is now the treatment of choice for subtotally resected and local central neurocytoma recurrence with higher tumour control rates and low complications despite the small number of patients studied in the literature.

Keywords: Central neurocytoma – Gamma knife surgery– stereotactic radiosurgery.


1. INTRODUCTION

Central neurocytoma (CN) is a rare brain tumor. They are typically located in the lateral ventricle near the foramen of Monro[1]. And are classified as World Health Organization (WHO) grade 2. These tumors were first described as a distinct tumor entity by Hassoun et al. in 1982 [2]. CN constitute 0.1–0.5% of all primary brain tumors [3, 4, 5, 6]. They are typically found in adolescents and young adults; with a higher incidence in the third decade, followed by the second and the fourth decades [7, 8, 9]. The male-to-female ratio is 1. [9, 10, 11, 12].

They are typically intraventricular and can occur hydrocephalus. Surgical removal is the standard treatment but, complete removal of the lesion is often difficult because of the deep location [13]. A combination of treatments, such as surgery with adjuvant radiation, can be considered for these lesions [14, 15]. A few number of studies have been published. Some authors have concluded that radiotherapy (RT) or stereotactic radio surgery can successfully control any residual tumour after incomplete resection or recurrence [16, 17, 18, 19, 20, 21].
The surrounding structures are therefore exposed to limited amounts of radiation. High-dose radiation for local lesions is possible with radio surgery. There are few reports on the efficacy of GKS for treatment of CNs. These reports are limited by the small number of patients studied [22, 23, 24, 25]. So, our study focuses on the efficiency of Gamma Knife radio surgery (GKS) in CN’s treatment.

2. Materials and Methods

2.1. Patients

We are relating the results of 11 patients who underwent GKS for CNs in the Gamma unit at functional and stereotactic neurosurgery service in Marseille from January 1999 to December 2015. One patient was excluded because the follow-up data was unavailable. The study included 4 men and 6 women. Their mean age at GKS was 25.12 years (range 17-41 years). The initial symptoms were headache in 10 patients, hemiparesis in 2 cases, gait disturbance in 3 cases, consciousness disturbance in 2 cases and ocular symptoms were found in 3 patients. All patients underwent surgery before GKS with a mean delay of 19.37 months (6-62 months). The resection was total in 2 patients, subtotal in 7 patients and one patient underwent only a biopsy. The histologic diagnosis was CN in all patients, and no atypical findings were noted. Seven (7) patients had a dilatation of ventricles. A ventriculo-peritoneal shunt was performed for hydrocephalus in only one patient.

2.2. Treatment

The procedure for GKS was as follows: a Leksell G stereotactic frame was attached to the patient’s head under local anesthesia, and a stereotactic magnetic resonance imaging (MRI) was performed. The treatments were planned with the Leksell gamma plan by the neurosurgeons. After confirming the tumour margin, an isodose curve fit to the tumour margin was made using 4 sizes of collimators. Finally, the prescribed mean dose delivered to the tumour margin was 19.1 Gy (12-24 Gy) with an isodose of 50%. The mean volume of treatment was 4.4 cm³ (1.03-10.7 cm³). Treatments were carried out according to the dose plan of the Leksell gamma knife model B, model C, model AC and Icon.

The follow up consists of a serie of cerebral MRI done at 6 months post GKS, then, once a year for 5 years and the 7th and 10th years post-GKS. The procedure was successful when there was a stabilisation of the lesion, a diminution of the tumour size or the involution of the tumour on the follow up MRI done at 3 years post GKS or 5 years post-GKS.

The decision of GKS treatment was made by a multidisciplinary team which include: neuroradiologists, radiation therapists and neurosurgeons.

2.3. Statistical Analysis

All factors that may affect the obliteration of the malformation, bleeding after the procedure and the occurrence of complications have been analysed with the statistical software SPSS 10 of Windows. Parameters with p < 0.05 were statically significant.

3. Results

On our study, 1 patient was lost from sight after the first control at 6 months, so we do not have his follow-up details. Of the remaining 10 patients, the mean time was 55.2 months (7.2-108 months). No deaths were observed during our follow-up.

3.1. Tumor Control

We had a volume reduction of all lesions treated, with an average of 86% (54-100%). Tumors were controlled locally in 7 cases including 3 cases who had a complete response. 3 patients had a partial response, and 1 case showed no change. The 3 and 5 year tumour’s control were 100%, (Fig 1). One recurring tumour was local (4.67 years after GKS) and 2 were “out-of-field” recurrence (5.42 and 6.8 years after GKS).

Figure1: A 17-year-old patient undergoing a central neurocytoma in 2009 with a ventriculo-peritoneal shunt preceding the radio surgical treatment. Here are the images illustrating the tumor control five years later.
A second treatment was performed in two patients with an average delay between the two of 5.91years. An average dose of 14.34Gy was performed and we had tumour control at 3 years of follow-up after the second GKS. In our follow-up, we had total lesional control in 9 patients (90%) during the whole follow-up period. The patient with local recurrence is still under surveillance.

3.2. Evaluation of the Patients
During our follow-up, there were no deaths in our series. The radio surgical treatment stabilized the lesions in nine patients. No patients underwent a second surgery. A VP shunt was performed for obstructive hydrocephalus at 12 months of radio surgery treatment. The treatment resulted in the disappearance of symptoms in almost all patients except for the two patients who had an hemiplegia, probably linked to surgical trauma. The Karnofsky Performance Scale Score remain stable after treatment.

3.3. Complications
We did not find in our study an intra-lesional haemorrhagic complication as reported in the literature. But we had symptomatic side effects of the radiation; a case of hydrocephalus by foramen Monro obstruction which required a VP shunt at 12 months of treatment. Even if it cannot be completely attributed to the rays; because even the tumour can obstruct the channels of the CSF.

4. DISCUSSION
CNs represent approximately 0.1-0.5% of all primary brain tumours. [5, 6]. Hassoun et al. described the first cases, since then, more than 500 cases have been described. These tumours typically affect adolescents and young adults with the highest incidence in the third decade, followed by the second and fourth decades [7, 8, 9]; it corroborates our study with a mean age of 25.12years (17-41years). The CN has been found in the cerebral hemispheres, spinal cord, brainstem, thalamus, amygdala, pineal gland, retina and cerebellum [2-8, 27, 28].

The treatment of choice for CN is complete resection of the tumour, usually leading to cure and long-term survival. However, in more than a half cases, the lesion cannot be completely resected [14, 15]. Gamma knife radio surgery is the treatment of choice for sub-totally resected and local CN recurrence [24, 25]. It is safer compared to conventional radiotherapy which can produce cognitive dysfunction and secondary tumour formation [20]. Although not statistically significant, Patel et al. [11] reported that adjuvant GKS for patients with STRs demonstrated 100% tumour control rate compared to patients without adjuvant fractionating radiation therapy (87% tumour control rate) [23]. Garcia et al. [20] also reported a higher tumour control rate of 93% with GKS versus 88% with fractionated radiation therapy.

Schild et al. [23] related the first cohort of GKS for CNs, the local control rate of CNs after radio surgery is reported to be 91% and 100% with 15Gy of marginal dose. Cobery et al. [29] rewired 4 patients with sub-totally resected or recurrent CN treated with GKS with a follow-up periods ranging from 12 to 99 months. They found an important reduction in tumour size, between 48% and 81%. Bertalanffy et al. [30] Anderson et al.[31] and Tyler-Kabara et al.[32] also reported similar results with a series of 3 to 4 residual or recurrent CNs, they found that patients had relatively excellent responses to GKS (reduction in tumour size, without complications). Karlsson et al. [25] reviewed 42 patients with CNs treated with GKS. The median prescribed dose was 13 Gy (range, 11-25 Gy), and the median tumour volume was 8 mL. They found that the local tumour control rate was unrelated to the prescribed dose (≤ 13 vs> 13 Gy, p=0.95) and was respectively 91% and 81% at 5 and 10 years after GKS. The local tumour control rate was also unrelated to tumour volume (< 7.55 vs > 7.55) cm3, p = 0.83).

Kim et al. [33] reported in a series of 13 patients treated by radio surgery in first intension had a volume reduction of 50% with no complications over a 6-96month follow-up period. Chen and al.[34] GKS to treat 14 large CNs with a mean volume of 19.6 mL and a mean prescribed dose of 12.1 Gy, 100% decreased in the 5years following GKS and a mean reduction rate of 69%. Yamanaka et al.[24] reported a median rate of tumour volume reduction of 74% (range, 15%-e100%). In the same report, the median prescribed dose was 15 Gy (range, 10-20 Gy), and the cumulative 5-year local control rate was 94%. On the basis of previous reports describing high local tumour control rates using a mean or median prescribed dose of 10.5-13 Gy. CN can be controlled with a relatively low prescribed dose. (See table).
In our series, the median prescribed dose and tumour volume were respectively 14.2 Gy and 4.4 mL. The cumulative local control rate was 90% at the last follow-up MRI. Our local tumour control was also unrelated to the prescribed dose (p= 0.89) and the tumour volume (p= 0.67) like in several reports [23, 24, 25].

We reported one local recurrence and 2 distant or out-of-field recurrence tumour years after radio surgery treatment. The last had a second session of radio surgery treatment for a local control, three years after the first procedure. Yamanaka et al. [24] reported, local recurrence in 2 cases (5.6%) after GKS. Karlsson et al. [25] reported 2 local recurrence (2.4 and 9.6 years after GKS) and 2 distant or out-of-field recurrences (1.6 and 4.8 years after GKS). Chen et al. [35] rewiered no recurrence after GKS in 14 CNs. Rades et al. [13] reported 1 recurrence in their 21 CNs, and Genc et al. [35] reported 1 recurrent case in their 22 CNs treated with GKS. The rate of local tumour recurrence is very low after GKS. Yen et al. [18] founded 2 out-of-field recurrences that were difficult to detect even with a retrospective review of imaging at the time of GKS in their 7 treated patients. Kim et al. [22] reported 3 out-of-field recurrences in their 20 CNs that were treated with GKS. All out-of-field recurrences occurred in cases undergoing surgery before GKS. Karlsson et al. [25] reported that a low incidence of distant recurrences does not justify a higher risk for long-term complications following radiotherapy. Local and distant tumour recurrences can be favourably managed using additional radiotherapy.

Yamanaka et al. [24] reported a radiation injury in 1 case and a slight memory disturbance persisted after the extent of the injury decreased. In this case, the tumour volume was 14.2 mL and the prescribed dose was 16 Gy. Martin et al. [36] founded 1 radiation necrosis case among their 4 CNs treated with linear accelerator radio surgery. The size of the tumour in this case was 23 mL, and the prescribed dose was 16 Gy. Chronic edema and radiation necrosis appeared 2 years after treatment, the patient’s symptoms improved after a ventriculoperitoneal shunt procedure and corticosteroid treatment [36]. Yen et al. [37] related a case that required a ventriculo-peritoneal shunt procedure due to hydrocephalus caused by an intratumoral hemorrhage 4 months after GKS. Gallina et al. [38] reported haemorrhage in the fourth ventricle and confirmed the absence of atypical histology with low proliferative potential after surgery. Terakawa et al. [39] also reported a hemorrhagic CN and speculated that approximately 3% of CNs was hemorrhagic. Karlsson et al. [25] reported no complications after GKS of 42 patients. In our study one patient developed hydrocephalus 12 months after GKS and required a ventriculo-peritoneal shunt. Yamanaka et al [24] reported that the incidence is low, a large lesion treated with a relatively high dose may tend to develop adverse radiation effects and the limit of the radio surgical dose for the fornix is not well known, symptoms associated with fornix injury may seldom occur with prescribed doses <15 Gy.

5. Conclusion

CN is a benign tumour of the CNS with an excellent prognosis. Surgery with gross complete resection is correlated with the best long-term survival rate and local tumour control. GKS is an effective treatment for residual and
recurrent CNs after surgery because of its high rate of local tumour control and the low morbidity.

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REFERENCES


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