

## Case Report

# Differential Diagnosis and Management of a Pituitary Mass with Renal Cell Carcinoma

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The small pituitary mass was incidentally found in 40-years-old women with renal cell carcinoma. The endocrinological and ophthalmological evaluation revealed no deficit and the short-term follow-up was recommended. In 6 months later, the visual disturbance was reported and the size of mass was increased. The tumor was removed totally via the trans-sphenoid approach. The post-operative endocrinological insufficiency was not noticed. During one year of follow-up period, there was no evidence of recurrence without adjuvant radiotherapy. The clinical features of pituitary metastasis from renal cell carcinoma were similar to those of pituitary adenoma. The possibility of pituitary metastasis should be kept in mind in patients with sellar mass and renal cell carcinoma.

**Key Words :** Pituitary · Metastasis · Renal cell carcinoma.

## INTRODUCTION

Metastatic tumors in the pituitary gland are uncommon and difficult to diagnose because most are small and clinically silent<sup>9</sup>). Only 6.8% of pituitary metastases are symptomatic, although a large autopsy series reported a 3.6% incidence of pituitary metastasis<sup>7,9</sup>). The original sites of malignancies in pituitary metastasis do not differ greatly from those in cerebral parenchymal metastasis. In pituitary metastasis, the most common primary malignancies are breast cancer in females and lung cancer in males, and other systemic metastases are often associated<sup>1,5</sup>). However, to the best of our knowledge, pituitary metastasis from renal cell carcinoma has been rarely reported, and only 25 cases have been published in the literature<sup>1,2,4</sup>). We present a case of pituitary metastasis from renal cell carcinoma.

## CASE REPORT

A 40-year-old woman presented with mild headache. Three years prior, she had undergone a left radical nephrectomy for renal cell carcinoma. Video-assisted thoracic surgery was performed for metastatic lymph node excision and left lower lobe

superior segmentectomy. Additional procedures included the excision of a retroperitoneal lymph node metastasis and thoracotomy for lymph node dissection along the descending aorta. She received chemotherapy for multiple metastases, including the adrenal gland, psoas muscle and mediastinum. During the follow-up after chemotherapy, she complained of intermittent headache. Magnetic resonance (MR) imaging demonstrated a 13 mm sellar mass with suprasellar extension without optic nerve compression (Fig. 1). The pituitary stalk was deviated to the left side. An endocrinological evaluation revealed hyperprolactinemia as follows : serum prolactin, 89.5 ng/mL [reference range (RR) : 0-25 ng/mL]; free T4, 0.75 ng/dL (RR : 0.70-1.80 ng/dL); TSH, 1.73 uIU/mL (RR : 0.4-4.1 uIU/m); ACTH at 8 am, 52 pg/mL (RR : 0-60 pg/mL); Cortisol at 8 am, 7.2 ug/dL (RR : 5-25 ug/dL); LH, 6.7 mIU (RR : 1-12 mIU); and FSH, 11.5 mIU (RR : 2-13 mIU). <sup>18</sup>F-fluorodeoxyglucose (FDG) positron emission tomography (PET) showed no uptake in the sellar area, and increased uptake in metastatic lesions in other sites (Fig. 1). She did not complain of the visual symptoms, and Goldmann perimetry revealed no visual field defects. A careful history revealed that she had experienced galactorrhoea and amenorrhoea, which continued for 5 months and resolved with-

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out management 9 years ago prior to her current presentation. With these clinical backgrounds, a presumptive diagnosis of pituitary adenoma was made and regular follow-up was recommended because of the relatively rare incidence of pituitary metastasis.

Six months after the first visit, her vision worsened. Snellen visual acuity without correction at a 5 m distance in 200 lux of light showed a marked decrease in visual acuity from 1.0/1.0 to 1.0/0.4. Goldmann perimetry revealed superior temporal quadrantanopsia of the left eye. The follow-up MR imaging showed that the mass from 13 mm to 23 mm, and compression of the optic apparatus was observed (Fig. 2). The cocktail test showed hyperprolactinemia (prolactin, 74.4 ng/mL) and hypocortisolism (peak cortisol, 11.9 ug/dL). The other pituitary axes were intact, and there were no symptoms suggesting diabetes insipidus.

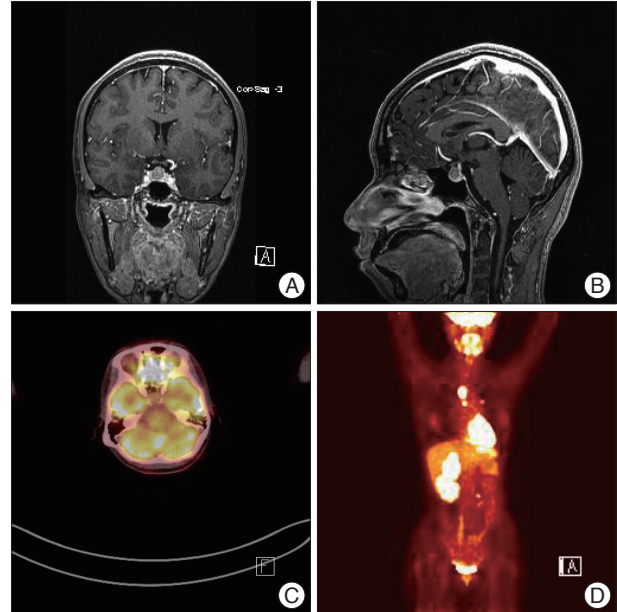
The patient underwent trans-sphenoid surgery to obtain a pathologic diagnosis and decompress the optic apparatus. The sellar floor was very thin and showed a mottled appearance. The tumor was gray and hypervascular (Fig. 3). Its hard consistency and the distinct demarcation helped to easily dissect it from the pituitary gland and cavernous sinus. However, we removed parts of the diaphragma sellae with the superior surface of the tumor because the tumor was strongly adhered to it. The tumor was completely removed in a piecemeal fashion, and the pathology was compatible with renal cell carcinoma (Fig. 4). The post-operative course was uneventful and her vision recovered to its previous original state within one month after surgery. The clinical and radiological follow-up has been continued without adjuvant therapy and hormone replacement.

## DISCUSSION

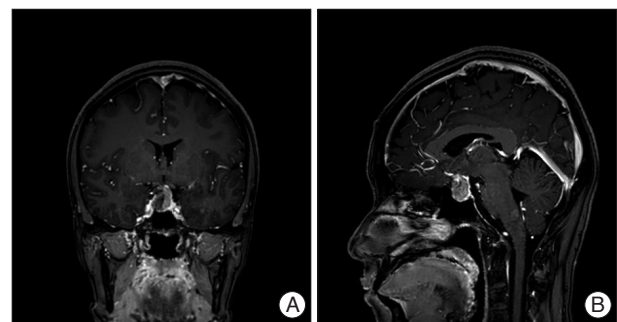
The pituitary gland is an uncommon site for the metastasis of systemic malignancies. Metastatic pituitary tumors constitute less than 1% of all surgical specimens from transsphenoidal surgery for sellar or parasellar tumors. Breast and lung cancer account for two thirds of pituitary metastases, although metastases from almost all cancers have been reported<sup>7,8</sup>. Additionally, renal cell carcinoma is a primary malignancy in only 2.6% of pituitary metastases<sup>9</sup>.

The differential diagnosis of pituitary metastases from other pathologies is challenging. Most metastatic pituitary tumors in-

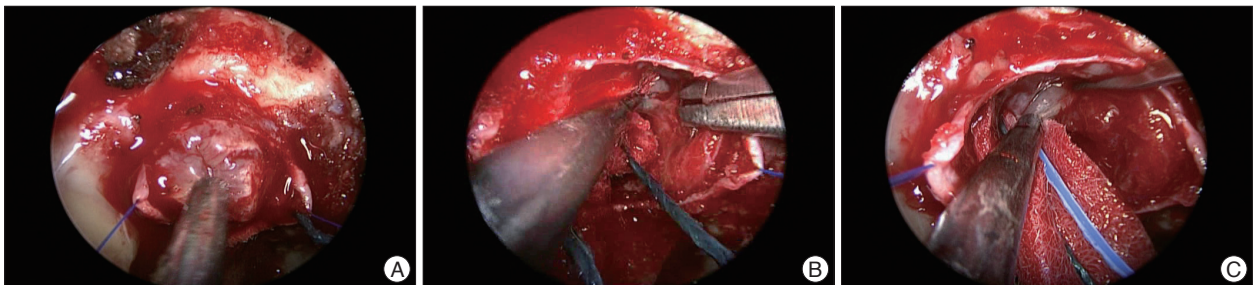
volve the posterior lobe<sup>6</sup>. A series of 190 symptomatic pituitary metastases identified the diabetes insipidus in 45.2% of patients<sup>4</sup>. The putative causes for frequent involvement of the posterior lobe were the relatively wide contact with the adjacent dura and



**Fig. 1.** The T1-weighted coronal (A) and sagittal (B) magnetic resonance images used to evaluate the intermittent headache reveal a round, well-enhanced sellar mass. The pituitary stalk is slightly deviated to the left by the tumor, and no optic compression is noted. 18F-fluorodeoxyglucose positron emission tomography shows no increased uptake in the sellar mass (C) in contrast to the other metastatic lesions in the body (D).

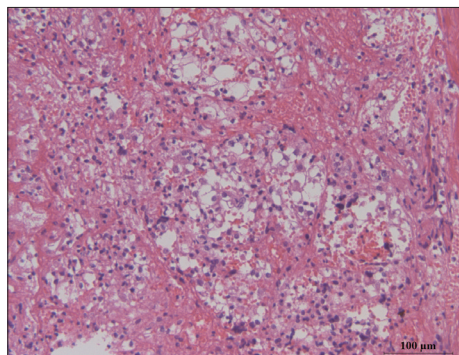


**Fig. 2.** The T1-weighted coronal (A) and sagittal (B) magnetic resonance images taken 6 months later show the markedly growing sellar mass compressing the optic chiasm.



**Fig. 3.** The tumor was gray and moderately hard (A). It has a sharp margin between the pituitary gland (B) and the diaphragma sellae (C). The dissection plane between the tumor and the surrounding normal structures is not strongly adhesive.

the direct arterial supply from the hypophyseal arteries, contrasting with the anterior lobe from the hypophyseal portal system<sup>6</sup>. Therefore, the presence of diabetes insipidus or cavernous cranial neuropathy in patients with primary malignancy suggests to pituitary metastasis over relatively common pathologies such as adenoma and Rathke's cleft cyst. However, pituitary metastasis from renal cell carcinoma shows similar symptoms and signs to pituitary adenoma. A recent study has revealed that the anterior pituitary dysfunction is more prevalent (90%) and diabetes insipidus is less prevalent (24%) in patients with pituitary metastases from renal cell carcinoma compared to those with other metastatic tumors<sup>4</sup>. These clinical features make it difficult to differentiate the pituitary metastasis from renal cell carcinoma from common pituitary pathologies with clinical features except the rapid growth rate as the presented case. Additionally, visual field defects are more commonly found in patients with renal cell metastases (82%) than those with other primary malignancies. Therefore, the trans-sphenoid surgery was recommended as the first-line treatment in all cases in literatures and stereotactic radiotherapy was performed in patients refused the surgery<sup>1,2,4</sup>. A half of patients underwent the radiotherapy as the adjuvant treatment (Table 1).



**Fig. 4.** The pathologic examination shows a nest of epithelial cells with clear cytoplasm and distinct cell membranes. The nests are separated by the highly vascularized stroma. Hematoxylin and eosin staining (×200).

In the reported case, we attempted to characterize a pituitary lesion with <sup>18</sup>F-FDG PET at the time of presentation. However, the lesion did not show uptake in <sup>18</sup>F-FDG PET, contrasting with the metastatic tumors in other sites, such as the adrenal gland and abdominal lymph nodes. The follow-up <sup>18</sup>F-FDG PET,

**Table 1.** Clinical features of the reported cases of pituitary metastasis from renal cell carcinoma

Sex	Age (years)	Interval from primary Dx. (years)	Status at diagnosis			Treatment	
			Hypopituitarism	Diabetes insipidus	Visual symptom	Primary	Adjuvant
Male	59	9	(+)	(-)	(+)	TSS	(-)
Male	53	0	(+)	(+)	(+)	TSS	RT
Male	75	9	(-)	(-)	(+)	TSS	(-)
Male	66	0	(+)	(-)	(-)	TSS	RT
Male	51	0	(+)	(-)	(+)	TSS	RT
Female	35	0	(+)	(-)	(+)	TSS	RT
Male	63	4	(+)	(-)	(+)	TSS	RT
Male	57	0	(+)	(+)	(+)	TSS	RT
Male	59	0	(+)	(+)	(+)	TSS	RT
Female	63	4	NA	NA	NA	TSS	(-)
Male	56	0	(+)	(-)	(-)	TSS	RT
Male	54	3	(+)	(-)	(+)	TSS	(-)
Male	73	8	(+)	(-)	(-)	TSS	(-)
Male	62	4	(+)	(+)	(+)	TSS	(-)
Female	77	0.25	(+)	(-)	(+)	TSS	SRT
Male	63	8	(+)	(+)	(+)	TSS	SRT
Male	70	6	NA	(-)	(+)	TSS	RT
Male	54	NA	(+)	(-)	(+)	TSS	RT
Male	67	27	(+)	(+)	(+)	TSS	RT
Male	51	12	(+)	(-)	(+)	TSS	RT
Male	53	0	(+)	(+)	(+)	TSS	RT
Female	67	11	(+)	(-)	(-)	SRT	Chemotherapy
Female	61	1	(+)	(-)	(-)	SRT	Chemotherapy
Male	74	5	(+)	(-)	(+)	TSS	(-)
Male	45	0	(+)	(-)	(+)	TSS	SRT
Female	40	3	(+)	(-)	(+)	TSS	(-)

TSS : trans-sphenoid surgery, RT : radiotherapy, SRT : stereotactic radiotherapy, NA : not available

which was performed 2 months before the vision disturbance, also did not show increased uptake in the pituitary gland. Retrospectively,  $^{18}\text{F}$ -FDG PET was not helpful for differentiating the pituitary metastasis from other pathologies in the reported case. Recently, a report revealed that various pathological abnormalities were identified using MR imaging in 52.7% of patients who showed focal  $^{18}\text{F}$ -FDG accumulation<sup>3</sup>). Additionally, the most common entity was pituitary adenoma, followed in incidence by metastasis and lymphoma. However, there was no significant difference in standardized uptake values between malignant and benign lesions. Therefore, the specific diagnosis of a pituitary lesion with  $^{18}\text{F}$ -FDG is not yet possible.

There were some characteristic of pituitary metastasis on the MR imaging, such as high signal intensity on T2-weighted images with iso- or hypointense signals, loss of high signal intensity in the posterior lobe on T1-weighted images, and a rapidly growing sellar mass with infiltrating features. However, these findings cannot be the pathognomic criteria for the diagnosis of pituitary metastasis, with the exception of growth rate. The rapid development of the visual field defect and growth on the follow-up MR images provided the clues for diagnosis. Therefore, the follow-up interval should be shorter in patients with asymptomatic pituitary lesions and systemic malignancies.

## CONCLUSION

The differential diagnosis of pituitary metastasis from a renal cell carcinoma from pituitary adenoma is challenging. Short-

term regular follow-up is mandatory in cases of pituitary lesions with underlying renal cell carcinoma.

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