

Punith Kempegowda<sup>1,2</sup>, Martyn Carey<sup>3</sup>, Latha Senthil<sup>4</sup>, Andrew Toogood<sup>1</sup> and John Ayuk<sup>1</sup>

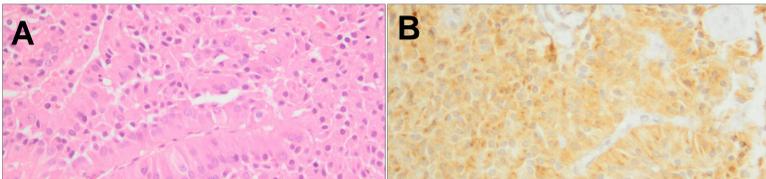
<sup>1</sup> Department of Endocrinology, Queen Elizabeth hospital Birmingham; <sup>2</sup> Institute of Metabolism and Systems Research, University of Birmingham; <sup>3</sup> Department of Neuropathology, Queen Elizabeth hospital Birmingham; <sup>4</sup> Department of Neuroradiology, Queen Elizabeth hospital Birmingham

## History and Examination

- A 61-year-old Caucasian male
- Presented with frequent hot flushes, dizziness, tinnitus and headache.
- No change in libido or shaving pattern.
- CT scan showed an incidental pituitary adenoma.
- Was diagnosed with essential hypertension when he presented with similar symptoms 2.5 years ago.
- No other co-morbidities and no family history of endocrine disorders.
- Clinical examination was essentially unremarkable.
- Intact visual fields.
- Testicles were normal in size for age.

## Investigations

- Unusually high follicular stimulating hormone (FSH) (>200 IU/L)
- Moderately raised prolactin (700 mU/L).
- Serum testosterone was marginally low (7.3nmol/L).
- MRI scan- large pituitary adenoma with right cavernous sinus involvement (Figure 1A).



**Figure 2: Histological images of pituitary tumour.** Haematoxylin and Eosin stained image (A) showing perivascular pseudo-rosette formation and Immunocytochemistry (B) image showing positive staining for FSH.

## Discussion

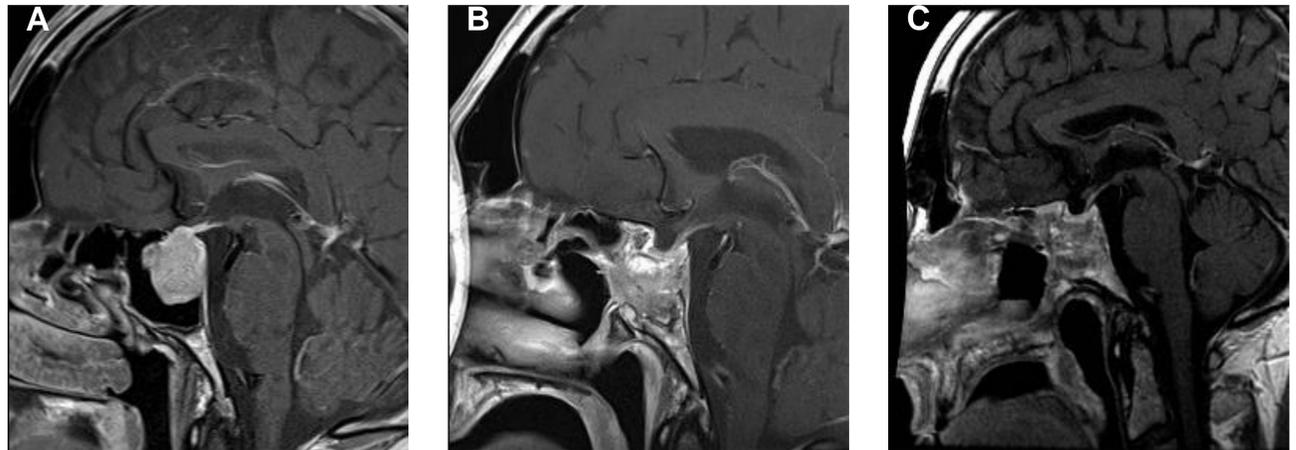
- Functioning gonadotrophic adenoma (FGA) represent a rare clinical entity causing distinct manifestations.
- They are similar in morphology to non-FGA.
- The first case related to a histologically confirmed FGA was reported in 1995 by Djerassi et al<sup>2</sup>.
- The most common clinical presentation include
  - Menstrual irregularity and ovarian hyperstimulation syndrome in females
  - Testicular enlargement in males
  - Isosexual precocious puberty in male children)
- The pathogenesis of FGA is unclear. Some of the proposed mechanism include<sup>1</sup>
  - Differences in GnRH receptor gene
  - FSH with an increased bioactivity to immunoreactivity ratio
  - Higher basic FSH isoforms are some of the proposed mechanisms.
- Figure 3 gives a simple algorithm to diagnose and manage patients with FGA.
- Treatment includes surgery, medical and radiotherapy.
- Life-threatening adverse effects such as pituitary apoplexy can rarely occur<sup>3</sup>.

## Summary

- We present a rare case of FGA in a man with successful resolution following combined surgical and medical treatment.
- Increased awareness is necessary for early diagnosis aiming to ameliorate the sequelae of hormonal hypersecretion and to minimize the consequences of the mass effect.

## References

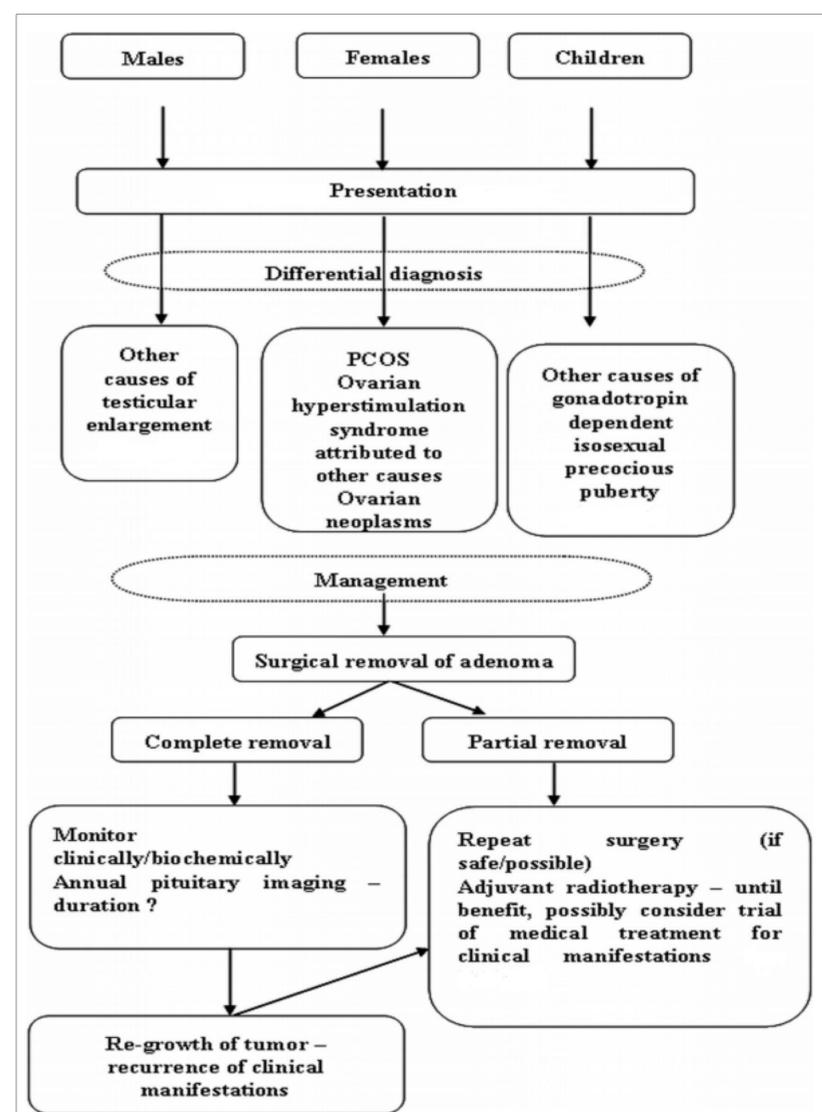
1. Ntali, G., Capatina, C., Grossman, A. & Karavitaki, N. Clinical review: Functioning gonadotroph adenomas. *J. Clin. Endocrinol. Metab.* **99**, 4423–33 (2014).
2. Djerassi, A. et al. Gonadotroph adenoma in a premenopausal woman secreting follicle-stimulating hormone and causing ovarian hyperstimulation. *J. Clin. Endocrinol. Metab.* **80**, 591–4 (1995).
3. Chanson, P. & Schaison, G. Pituitary apoplexy caused by GnRH-agonist treatment revealing gonadotroph adenoma. *J. Clin. Endocrinol. Metab.* **80**, 2267–2268 (1995).



**Figure 1: Magnetic Resonance Imaging (MRI) pituitary- pre-op (A)** showing a large tumour of the pituitary fossa with suprasellar extension in contact with the chiasm and suprasellar nodules up the pituitary stalk, **Post-op (B)** demonstrating a reduction in tumour burden indicated by the resolution of pituitary stalk to near normal anatomical position; however the solid tumour indicating a significant residual disease **and post-octreotide treatment (C)** shows reduction in secretion of FSH indicated by its cystic nature.

## Management

- Elective trans-sphenoidal debulking resection of adenoma in view of size and uncertainty of the nature of lesion.
- Histopathology- FSH secreting pituitary adenoma (Figure 2).
- Residual adenoma despite the surgery (Figure 1B) and his FSH remained high (187.5 IU/L) post-surgery.
- Trialled with octreotide- FSH levels normalised (11.2 IU/L) and his residual tumour shrunk (Figure 1C).
- Currently, patient continues to enjoy the period of symptom resolution and good health.



**Figure 3: Flowchart with differential diagnosis and management of patients with FGA.** From: Ntali et al.<sup>1</sup>

Queries to: p.kempegowda@nhs.net