





# Acromegaly caused by atypical pituitary adenoma

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#### INTRODUCTION

- Diagnostic criteria for an atypical adenoma include invasive growth, elevated mitotic index, Ki-67 labeling index greater than 3% and extensive p53 immunoreactivity.
- Atypical pituitary adenomas have higher risk of aggressive behavior in particular by the higher growth, local invasion and high risk of recurrence after surgery.
- In acromegaly the resistance to somatostatin analogues may be another manifestation of these adenomas since higher levels of Ki-67 are associated with poor response to therapy.
- Radiotherapy is reserved for when surgery and medical therapy fails to control the disease. The experience of the use of temozolomide in aggressive adenomas is extremely limited.

## **CASE REPORT**

#### Identification

41 year-old, woman

#### Past medical history

- Carpal tunnel syndrome
- Thyroid nodule
- Arthralgia

**Therapy**: trazodone 100mg/day, bromazepam 6mg/day

# History

- Headaches with 8 years of evolution
- Growth of the hands
- Hyperhidrosis
- Enlargement of the lips with 3 years of evolution
- Episode of visual change that reversed spontaneously

Was referred to the Endocrine team

#### Physical examination

- W= 66.5 Kg, H= 1.57m, BMI= 27Kg/m2
- Visual field examination normal

### Exam already carried out

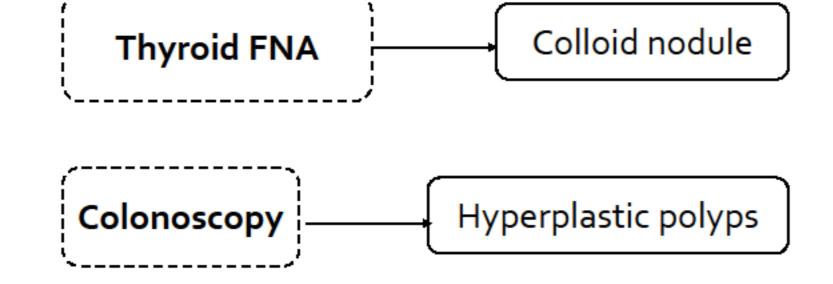
 Pituitary CT: Pituitary mass lesion hourglass-shaped. The lesion appears to invade the tank Meckel

Table 1. Hormonal study						
Parameter	Result	Reference range				
GH	25.6ng/mL	<8				
IGF-1	1689ng/mL	64-336				
PTH	50.2pg/mL	10.0-65.0				
T <sub>4</sub> L	0.95 ng/dL	0.70-1.48				
TSH	0.88uUI/mL	0.35-4.94				
Prolactin	21.2 ng/mL	4.8-23.3				
Cortisol	10.9 ug/dL	6.2-19.4				

Oral Glucose Tolerance Test	st	Time (min)	Glucose (mg/dL)	Insulin (uU/mL)	GH (ng/mL)
	-	0	106	32.6	24.1
	nce	30′	199	186.2	17.3
	lera	6o'	197	199.8	15.8
	<b>1</b> 0	90′	207	330.8	14.7
		120	204	431.9	12.7

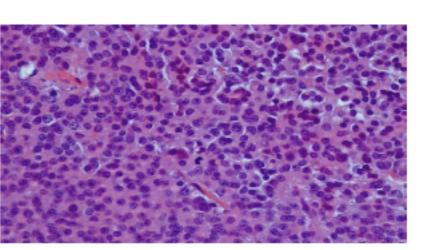
#### Pituitary MRI – december 2013

Pituitary lesion with 24mm craniocaudal diameter and 5 mm transverse diameter with invasion of the cavernous sinus.



She was submitted to transphenoidal resection in august 2014

## Clinical pathology report:



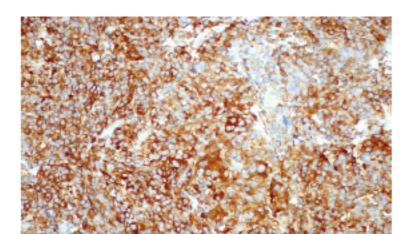
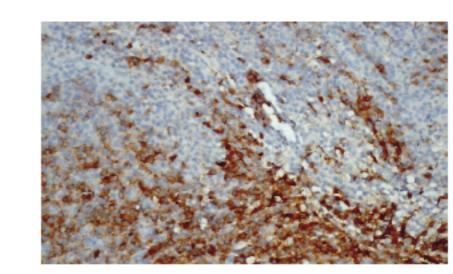


Fig.1 – Presence of mitosis

Fig.2 – GH: diffuse expression



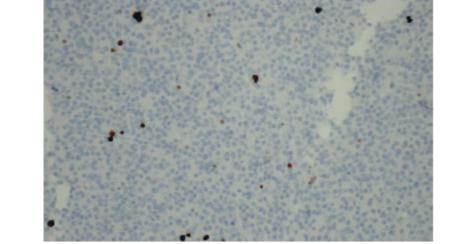
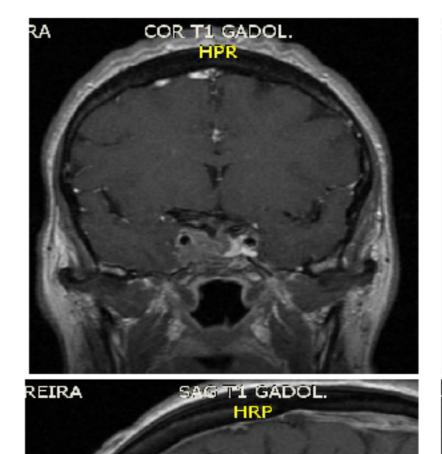
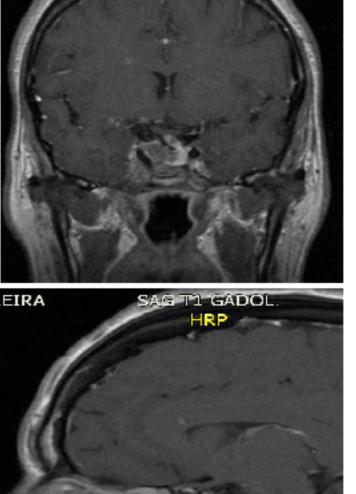


Fig.3 – PRL: multifocal expression

Fig.4 – **Ki67** labelling index was **8%** 





There is no longer compression of the optic chiasm. Persistence of tissue attributable to residual tumor invading the right cavernous sinus.

Fig. 5 - Pituitary MRI november 2014

Table 2. Oral Glucose Tolerance Test –							
november 2014							
Tempo	Glucose		GH				
(min)	(mg/dL)	(uU/mL)	(ng/mL)				
0	86	32.8	21.8				
30	129	80.7	17.4				
60	171	75-5	13.9				
90	154	217.5	12.8				
120	126	188.6	11.1				
-	-	-	-				

The patient began treatment with somatostatin analogs - octreotide

## CONCLUSIONS

➤ This is a case of acromegaly with several predictors of a possible failure of response to somatostatin analogues: young age, elevated levels of GH and IGF-1 at diagnosis, macroadenoma with invasive growth and Ki-67 labeling index of 8%. In the presence of cases of atypical adenomas a multidisciplinary team of endocrinologists, neurosurgeons, pathologists and oncologists must work together in attentive and personalized follow-up of the patient in order to prevent disease progression.

References: Katznelson L, Laws ER Jr, Melmed S, Molitch ME, Murad MH, Utz A, Wass JA. Acromegaly: and endocrine society clinical practice guidelines. J Clin Endocrinol Metab. 2014 Nov;99(11):3933-51; Carracasco CA, Gadelha M, Manavela M, Bruno OD. Agressive tumors and difficult choices in acromegaly. Pituitary. 2014 Jan;17 Suppl 1:S24-9









