# Efficacy & predictive factors of response to immunotherapy in pituitary carcinomas & aggressive pituitary tumors: a French cohort study

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**CONTEXT:** After temozolomide failure, no evidence-based treatment is available for pituitary carcinomas (PCs) & aggressive pituitary tumors (APTs). So far, only 12 cases treated with immune-checkpoint inhibitors (ICIs) have been published, showing encouraging efficacy. Predictive factors of response are lacking.

**OBJECTIVE:** assess the real-life efficacy and predictors of response to ICIs in PCs and APTs.

#### **MATERIAL & METHODS**

- > Multicentric, retrospective, observational cohort study, including all PCs and APTs treated with ICIs in France up to March 2022.
  - > PD-L1 immunohistochemistry and CD8+ T cell infiltration were evaluated centrally.

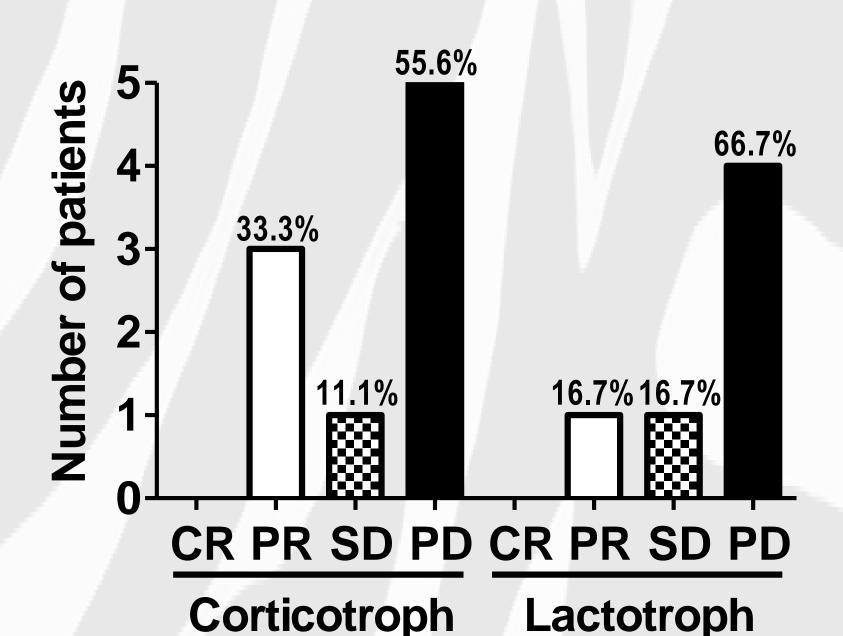
#### I. Treated cases

#### Corticotroph Lactotroph 6 cases 9 cases Sex Male 5 (55.6%) 5 (83.3%) Female 4 (44.4%) 1 (16.7%) 31-72 39-75 Age at treatment Tumor type 4 (44.4%) PC 2 (33.3%) **APT** 5 (55.6%) 4 (66.7%)

all patients had previously received multimodal treatment, including temozolomide

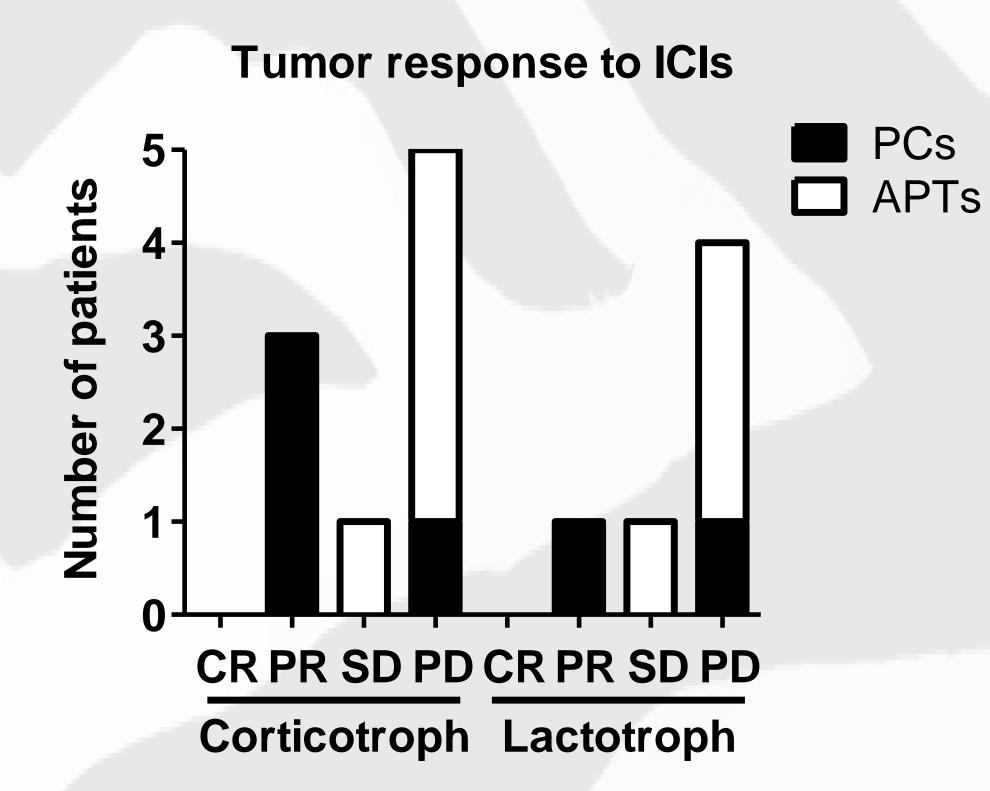
#### **RESULTS**

### II. Tumor response to ICIs



- ➤ Lower real-life efficacy of ICIs compared to previously published data.
- Slightly better tumor response of corticotroph compared to lactotroph cases.

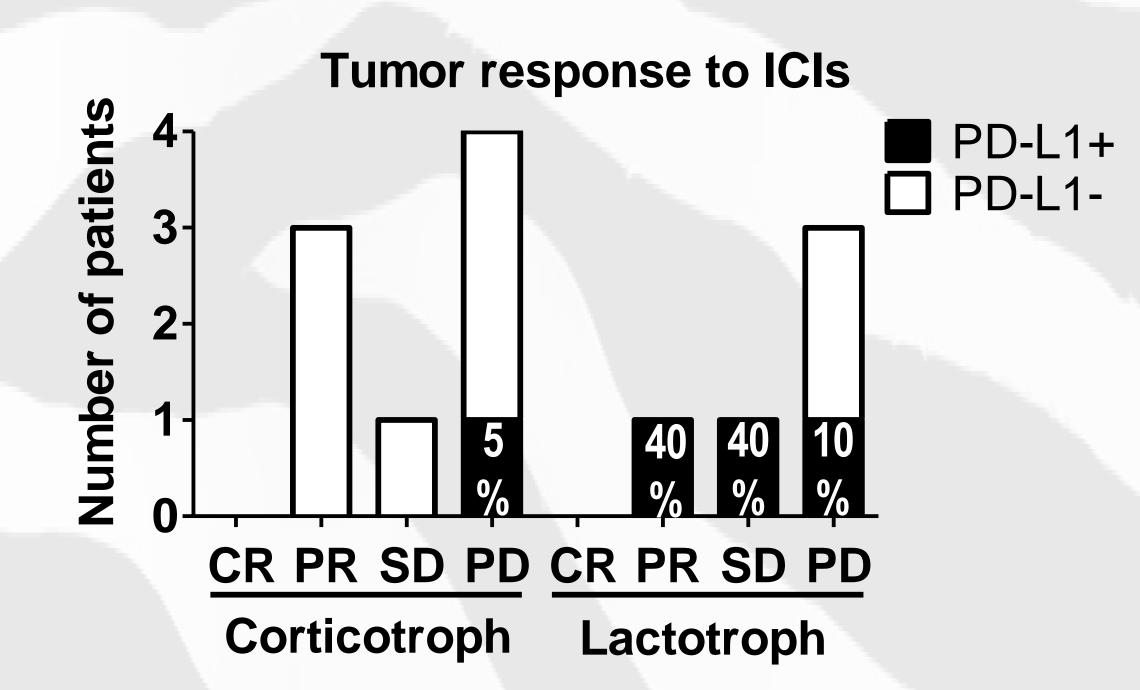
#### II. Predictive factors of response – tumor type (PC vs APT)

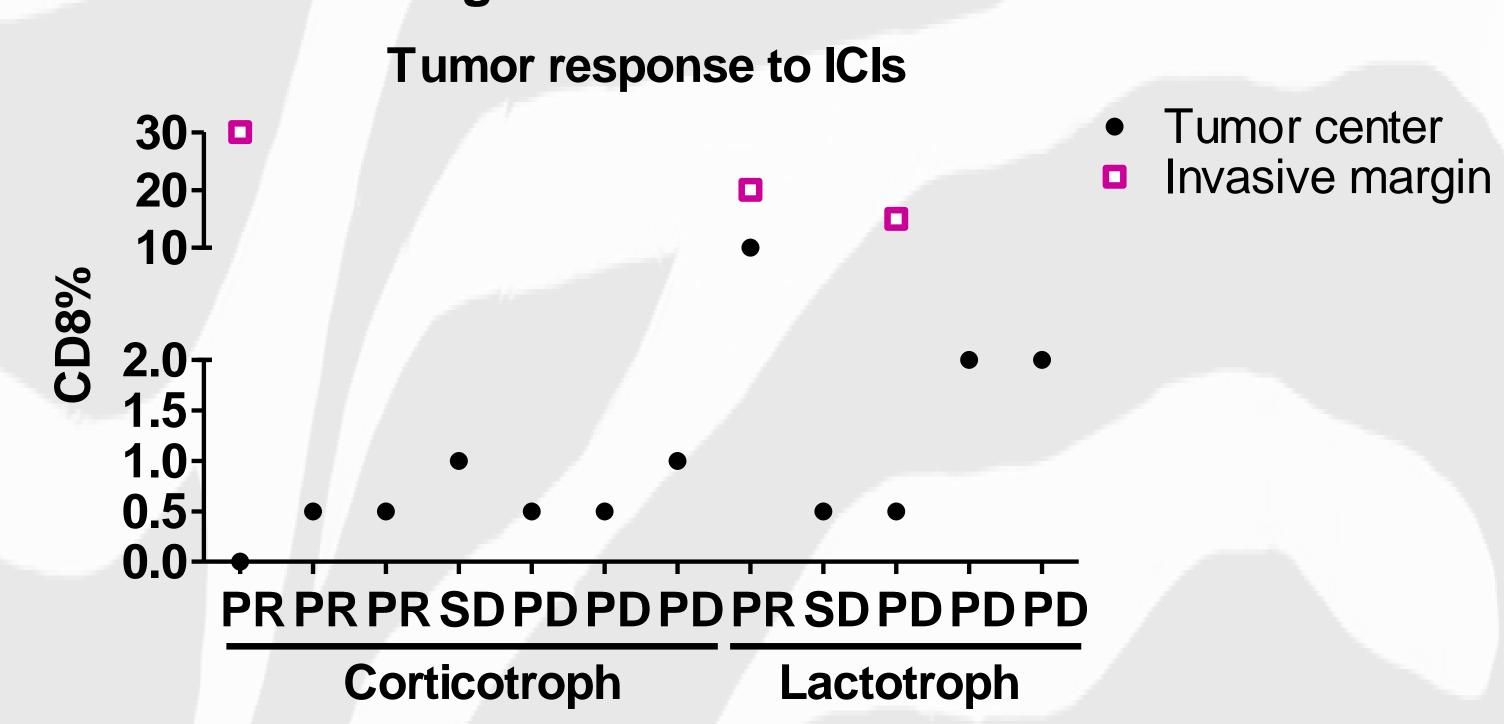


#### PCs responded far better than APTs!

- > An extension work-up is needed to confirm or exclude metastases before starting ICIs.
- > For PCs (especially corticotroph), ICIs are a reasonable 2<sup>nd</sup> line therapeutic option after temozolomide.
- ➤ Underlying biological differences between PCs and APTs?

#### III. Predictive factors of response – PD-L1 staining & CD8+ T cell infiltration





Negative PD-L1 staining & very low CD8+ T cell infiltration in the tumor center should not preclude ICI use in corticotroph PCs!

Further validation is warranted in lactotroph tumors.

## CONCLUSIONS

- 1 Tumor type (PC versus APT) is a major predictor of response to ICIs.
- 2 After temozolomide failure, ICIs should be proposed for PCs, especially for corticotroph PCs, being the best currently available option.
- 3 Negative PD-L1 staining & very low CD8+ T cell infiltration in the tumor center should not preclude ICI administration in corticotroph carcinomas.