Re: Oncocytoma on renal mass biopsy: is it still the same histology when surgery is performed? Results from UroCCR-104 study. N Branger et al https://doi.org/10.1007/s00345-022-04261-3

# Oncocytoma on renal mass biopsy: why is surgery even performed?

Hannah Warren, Carlotta Palumbo, Anna Caliò, Maxine GB Tran, Riccardo Campi; Collaborators§

On behalf of the European Association of Urology (EAU) Young Academic Urologists (YAU) Renal Cancer Working Group

### § Collaborators:

Jean Courcier, Zhenjie Wu, Riccardo Bertolo, Angela Pecoraro, Daniele Amparore, Leonardo D. Borregales, Chiara Ciccarese, Pietro Diana, Selcuk Erdem, Laura Marandino, Stijn Muselaers, Nicola Pavan, Umberto Carbonara, Eduard Roussel, Michele Marchioni

Contemporary management of localized renal masses (LRM) is nuanced, balancing oncologic control against overtreatment. For patients harbouring benign and asymptomatic oncocytoma, upfront partial or radical nephrectomy represents a diagnostic rather than therapeutic procedure, with risk of serious adverse events(1). Yet, the role of preoperative renal tumour biopsy remains controversial.

We welcome Branger and colleagues' contribution from the multicentre French UroCCR project (NCT03293563) (2), but we are puzzled by the high proportion of patients with biopsy-diagnosed oncocytoma managed with surgery (63%), either upfront or after a period of surveillance.

Biopsy-diagnosed oncocytomas on active surveillance (AS) exhibit low growth (1-2 mm/yr) with few transitioning to active treatment (3-11%) after median follow up of 29-34 months (3–5), with no treported metastatic events or deaths (6). The oncological safety of AS for all LRMs (benign and malignant) has also been reported at a mid-/long- term follow-up(7).

In contrast, Branger et al report that 35% of patients managed with initial AS transitioned to surgery during a median follow up of 28 months, on the basis of 'growth during surveillance' in 96% of cases. Median growth rate in this group would have been of interest but was not reported.

Surgical overtreatment of oncocytoma reflects the uncertainty regarding the positive predictive value of renal mass biopsy (RMB), particularly surrounding misclassification of histology/grade due to tumour heterogeneity(8,9). Of note, the vast majority of tumours "misclassified" at RMB in the study by Branger et al were chromophobe or other oncocytic/chromophobe renal cell carcinoma (RCC) on final pathology(2). While the latest WHO classification of urogenital tumours cautioned against a *definite* diagnosis of oncocytoma from a needle core biopsy(8), we join the authors in emphasizing that such misclassification might be of limited clinical significance, given the low metastatic potential and excellent prognosis of tumours on this spectrum(4). Reassuringly, clear cell and unclassified RCC were found in only 4% of operated cases.

Discordance between RMB and surgical pathology for samples obtained at centres inside and outside the French Research Network for Kidney Cancer was 21% vs 52%, respectively, calling into question the appropriateness of performing and reporting RMB outside high-volume institutions. A contemporary pragmatic objective for RMB would be to reliably differentiate "low grade oncocytic neoplasia" from other (more aggressive) RCC histotypes. Such discrimination may allow decision making with clinically meaningful impact.

Lastly, upfront surgery was performed most commonly due to 'tumour size' or 'surgeon decision', with only 3% of patients undergoing surgery based on patient preference alone; this suggests that patients accept AS when offered. A more detailed justification for the decision to operate and the rationale for biopsy in the first instance would have been desirable, since a benign result did not allow the patient to avoid surgery.

In conclusion, the study by Branger et al. highlights the ongoing challenge of reducing overtreatment of benign, often incidental renal tumours. No test is perfect, but judicious use of RMB and novel "virtual biopsy" imaging tools (10) will be key to risk-stratify LRMs, allowing objective and individualized counselling towards informed, evidence-based and shared decision-making.

### **Authors' Contribution**

Hannah Warren: Project development, manuscript writing Carlotta Palumbo: Project development, manuscript writing

Anna Caliò: Project development, manuscript writing

Maxine GB Tran: Manuscript editing

Riccardo Campi: Project development, manuscript writing

### **Collaborators' contribution**

Jean Courcier: Project development, manuscript editing

Leonardo D. Borregales: Project development, manuscript editing

Laura Marandino: Project development, manuscript editing

Zhenjie Wu: Project development
Riccardo Bertolo: Project development
Angela Pecoraro: Project development
Daniele Amparore: Project development
Chiara Ciccarese: Project development
Pietro Diana: Project development
Selcuk Erdem: Project development
Stijn Muselaers: Project development
Nicola Pavan: Project development

Umberto Carbonara: Project development Eduard Roussel: Project development Michele Marchioni: Project development

### Disclosure of potential conflicts of interest

The authors have relevant conflicts of interest to declare

# **Research involving Human Participants and/or Animals**

No human participants or animals were involved in the development of this manuscript

## **Informed consent**

Not Applicable

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