

## PRESS RELEASE

ASCO 2025 – Oral abstract session

Villejuif, 30 May 2025

### A COMBINATION THERAPY SHOWS SIGNS OF EFFICACY IN TREATMENT-RESISTANT PATIENTS WITH A SUBTYPE OF COLORECTAL CANCER

*Dr Antoine Hollebecque, oncologist in the Department of Therapeutic Innovation and Early Trials (DITEP) at Gustave Roussy, presents the results of a phase I/II trial at the ASCO 2025 Congress. The study evaluated a novel targeted combination therapy in patients with advanced colorectal cancer carrying the KRAS G12C mutation. It focused on olomorasib, a second-generation KRAS inhibitor, combined with cetuximab, a monoclonal antibody targeting the EGFR receptor. **This therapeutic approach aims to overcome resistance to standard treatments in heavily pretreated patients, often left with no remaining options.***

**Abstract no. 3507 presented orally by Dr Antoine Hollebecque on Friday 30 May at 16:57 UTC -5.**

This oral presentation is one of the 109 presentations featured in the 2025 ASCO programme with contributions from Gustave Roussy's physician-scientists. Gustave Roussy is active across numerous areas of expertise, reflecting both the high quality of the research carried out at the Institute and its international standing.

Colorectal cancers are among the most common in France, with over 47,000 new cases diagnosed in 2023<sup>1</sup>. The treatment of the metastatic disease largely relies on a combination of chemotherapy and targeted therapy. However, for patients with advanced tumours, therapeutic options remain limited, particularly after failure of standard chemotherapy regimens such as FOLFOX and FOLFIRI. The KRAS G12C mutation, found in approximately 3 to 4% of colorectal cancers, is associated with poor prognosis and resistance to EGFR inhibitors when used alone.

It is in this context that the international phase I/II LOXO-RAS study, sponsored by Loxo Oncology, has been conducted. LOXO-RAS aimed to evaluate the combination of a second-generation KRAS G12C inhibitor, olomorasib, with the anti-EGFR targeted therapy cetuximab. The goal? To overcome resistance mechanisms by combining two complementary approaches to improve treatment response in heavily pre-treated patients. The results have been presented at the ASCO 2025 Congress by Dr Antoine Hollebecque, oncologist at the Department of Therapeutic Innovation and Early Trials (DITEP) at Gustave Roussy.

#### A Promising Response Rate

The trial enrolled 93 patients with metastatic KRAS G12C-mutated colorectal cancer, who had received an average of three prior lines of therapy. Two doses of olomorasib were tested, and

<sup>1</sup> Institut national du cancer. *Panorama des cancers en France – Édition 2024*. INCa ; 2024. Available at : <https://www.e-cancer.fr/Expertises-et-publications/Catalogue-des-publications/Panorama-des-cancers-en-France-edition-2024>

the novel treatment was generally well tolerated, with most adverse events being grade 1 or 2, mainly gastrointestinal or dermatological – typical with an anti-EGFR combination.

Among these patients, often at the end of therapeutic options, the LOXO-RAS trial results were highly encouraging. The objective response rate — including complete and partial responses — reached 42%. Disease control was observed in over 90% of cases. Progression-free survival reached 7.4 months. These results are considered positive in such a difficult-to-treat population.

*“These data confirm the relevance of combination approaches to circumvent resistance in KRAS G12C-mutated cancers,”* says Dr Antoine Hollebecque. *“They pave the way for new personalised treatment strategies in situations where options have until now been extremely limited,”* he concludes.

This combination could eventually become part of the therapeutic arsenal for patients with advanced colorectal cancer carrying this specific mutation. The results of the LOXO-RAS study support the launch of phase III registration trials using these new-generation KRAS G12C inhibitors.

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#### Abstract no. 3507

*Efficacy and safety of olomorasib, a second-generation KRAS G12C inhibitor, plus cetuximab in KRAS G12C-mutant advanced colorectal cancer.*

*Oral abstract session presented by Dr Antoine Hollebecque.*

*Friday, 30 May 2025 | 16:57 UTC-5.*

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#### About Gustave Roussy

Ranked first in France, first in Europe and fourth in the world, Gustave Roussy is a centre of global expertise entirely dedicated to patients living with cancer. The Institute is a founding pillar of the Paris Saclay Cancer Cluster. Source of therapeutic innovations and diagnostic breakthroughs, the Institute welcomes nearly 50,000 patients each year, including 3,500 children and adolescents, and develops an integrated approach combining research, care and teaching. An expert in rare cancers and complex tumours, Gustave Roussy treats all cancers at all stages of life. It offers its patients personalised care that combines innovation and humanity, taking into account both care and the physical, psychological and social quality of life. With 4,100 employees at two sites, Villejuif and Chevilly-Larue, Gustave Roussy brings together the expertise essential for high-level cancer research; 40% of treated patients are included in clinical studies. To find out more about Gustave Roussy and follow the Institute's news: [www.gustaveroussy.en](http://www.gustaveroussy.en), [X](#), [Facebook](#), [LinkedIn](#), [Instagram](#) and [Bluesky](#).



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