Metastatic colorectal cancer: don't make things worse!

Abstract

- Despite treatment with the Folfox protocol followed by the Folfiri protocol (or the reverse sequence), about 50% of patients with metastatic colorectal cancer die within less than 2 years. *Cetuximab*, a monoclonal anti-EGFR antibody, has no proven impact on survival after failure of these protocols.
- Panitumumab (Vectibix°, Amgen) is another monoclonal anti-EGFR antibody authorised for use in this setting, but only

panitumumab

(VECTIBIX°)

Solution for IV infusion (after dilution)

- 100 mg panitumumab per vial (20mg/ml)
- 400 mg panitumumab per vial (20mg/ml)
- Licensed indication: "(...) monotherapy for the treatment of patients with EGFR expressing metastatic colorectal carcinoma with non-mutated (wild-type) KRAS after failure of fluoropyrimidine-, oxaliplatin-, and irinotecan-containing chemotherapy regimens".

[EU marketing authorisation, centralised procedure]

cytotoxic agent; monoclonal anti-EGFR antibody

when tumour cells express EGFR and bear the wild-type (unmutated) KRAS gene.

- The initial clinical evaluation only includes one randomised unblinded trial comparing *panitumumab* plus symptomatic treatment versus symptomatic treatment alone in 469 patients. About half of the patients died during the first 6 months of treatment, with no difference between the groups. The addition of *panitumumab* "significantly" increased the median progression-free survival time, but only by 5 days (8.0 weeks versus 7.3 weeks). A retrospective subgroup analysis suggested that *panitumumab* might be more useful in patients with the wild-type KRAS gene.
- As expected, given panitumumab's mechanism and its proteinaceous nature, its adverse effects are similar to those of cetuximab, with skin disorders in almost all patients (especially acne-like rash), as well as pulmonary disorders, diarrhoea, hypersensitivity reactions.
- Adding panitumumab to bevacizumab and/or to oxaliplatin- or irinotecan-based chemotherapy regimens leads to a major increase in its adverse effects.
- When the two most widely used protocols fail to control metastatic colorectal cancer, it is better to focus on maintaining patients' quality of survival, by means

of appropriate symptomatic treatment, rather than resort to *panitumumab*, which has more adverse effects than documented benefits.

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NOT ACCEPTABLE



Adding panitumumab to symptomatic treatment in patients with

metastatic colorectal cancer who failed to respond to the two most widely used protocols scarcely increases survival or progression-free survival. Like other monoclonal anti-EGFR antibodies, panitumumab provokes sometimes serious adverse effects in most patients: epithelial toxicity (skin) and hypersensitivity reactions.

Rev Prescrire 2008; 28 (301): 817.



In response to our request for information, Amgen provided us with published material only.

Full review (5 pages, 11 references) available in French on request *Rev Prescrire November 2008; 28 (301): 817.*

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