

Optimizing strategies for the treatment of colorectal cancer in the biomarker era



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### **Expert panel**



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## Agenda

**Evidence-based first-line treatment choices for patients with CRC** 

Implementing biomarker testing in the clinic: Practical considerations

Will novel combination regimens be the future of CRC treatment?



# **Evidence-based first-line treatment choices for patients with CRC**

#### Dr Kei Muro

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## Established and emerging biomarker-guided treatment options for mCRC

Biomarker	Targeted drugs	Resistance
RAS	Small molecules targeting G12C mutation	Confers anti-EGFR agent resistance
BRAF	BRAF inhibitors	Negative predictor of response to anti-EGFR therapies
Microsatellite instability	Pembrolizumab, nivolumab (PD-1 inhibitors), ipilimumab (CTLA4 inhibitor)	
HER2	Trastuzumab, lapatinib, pertuzumab, trastuzumab deruxtecan	Predict resistance to anti-EGFR therapies
NTRK	Entrectinib, larotrectinib	



BRAF, v-Raf murine sarcoma viral oncogene homolog B; CTLA4, cytotoxic T-lymphocyte-associated protein 4; EGFR, epidermal growth factor receptor; HER2, human epidermal growth factor receptor 2; mCRC, metastatic colorectal cancer; NTRK, neurotrophic tyrosine receptor kinase; PD-1, programmed cell death protein 1; RAS, rat sarcoma virus. Crutcher M, Waldman S. *Front Med.* 2022;9:1062423.

# Implementing biomarker testing in the clinic: Practical considerations

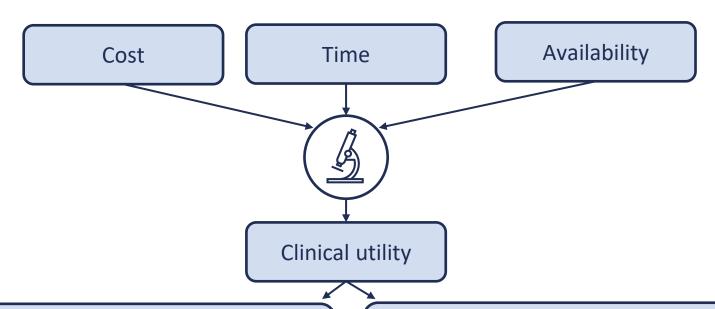
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### Practical considerations for biomarker testing



#### **Prognostic biomarkers**

Inform about likely disease outcome independent of the treatment received

#### **Predictive biomarkers**

Provide information about likely outcomes with application of specific interventions



# Will novel combination regimens be the future of CRC treatment?

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### **Combination therapies**

Many combination therapies for CRC using different biomarker analysis are under investigation including:

#### **KRAS G12C mutation**

- Sotorasib + panitumumab (NCT04185883)
- Sotorasib + bevacizumabawwb (NCT04185883)

#### **BRAF V600E mutation**

- Encorafenib + cetuximab ± binimetinib (NCT02928224)
- Encorafenib +
   binimetinib +
   cetuximab
   (NCT03693170)

#### MSI-H/dMMR

- Atezolizumab + bevacizumab (NCT02997228)
- Nivolumab + ipilimumab (NCT02060188)

#### **HER2** positive

- Trastuzumab
  emtansine +
  pertuzumab
  (NCT03225937)
- Trastuzumab + pyrotinib (NCT04380012)

