

## Colorectal cancer – *RAS* Mutation Testing on ctDNA

### Background

Mutations in the *KRAS* oncogene (OMIM 190070) and *NRAS* oncogene (OMIM 164790) are frequently found in a number of human cancers. Clinical data has established that 51% of metastatic colorectal tumours may carry a mutated *RAS* gene. An activating mutation indicates that small molecule drugs (such as Erlotinib and Gefitinib) or drugs which target the Epidermal Growth Factor Receptor (EGFR) (such as Cetuximab and Panitumumab) will be ineffective in terms of control of tumour proliferation.

The recent development of circulating tumour DNA (ctDNA) technologies has presented us with new opportunities to improve the stratification of colorectal patients for treatment. ctDNA is shed into the patient's blood stream from the tumour. The ctDNA is representative of the tumour and therefore presents a viable alternative to an invasive procedure for molecular analysis for *RAS* mutations.

### Recommended Clinical Referral Criteria

Patients with metastatic colorectal cancer, suitable for treatment with Cetuximab, where either no biopsy sample is possible, or where the biopsy sample has been exhausted or has failed molecular analysis.

### Molecular Analysis

**Mutation hotspot screen: Droplet digital PCR** analysis of the most common *RAS* mutations in *KRAS* codons 12, 13 and 61 and *NRAS* codon 61 (covers 96% of known *KRAS* mutations and 47% of known *NRAS* mutations in colorectal cancer). The analysis will detect ~0.5%-1% of mutant in a background of wild-type genomic DNA

### Prices\* & Turnaround Times (TAT)

\*Prices available on request

Test	TAT (calendar days)
Analysis for <i>KRAS</i> codons 12, 13 and 61 and <i>NRAS</i> codon 61	7

### Contact Details

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Accredited to ISO 15189:2012  
(8988)

### Sample Requirements

10ml blood in either CellSave© or Streck©  
tube to reach the laboratory within 24 hours;  
it is recommended that samples are not  
taken whilst patient is receiving  
chemotherapy.

Please label samples with three identifiers  
and date of collection.

**All samples must be accompanied by  
request form.**

Consent for testing & DNA storage is  
assumed when request for test received

Websites for tubes:

[www.streck.com](http://www.streck.com) [www.janssen.com](http://www.janssen.com)

### Links

Orphanet

<http://www.orpha.net/>

EDDNAL

<http://www.eddnal.com/>

OMIM

<http://www.omim.org/>

Genetic Test Registry

<http://www.ncbi.nlm.nih.gov/gtr/>

Support

Bowel Cancer UK

<http://www.bowelcanceruk.org.uk/>