Know Cancer's Next Move

- > Does my patient need adjuvant chemotherapy?
- > Is the treatment working?
- > Is the cancer recurring?

Discover unique insights with molecular residual disease (MRD) detection, informed by ctDNA

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Signatera™ Residual disease test (MRD)

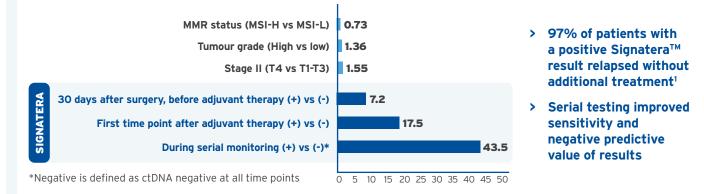
L'feLabs GENETOCS

When to use Signatera[™] for patients with CRC

In the adjuvant setting

Use after surgery to evaluate the need for adjuvant chemotherapy and potentially avoid unnecessary treatment

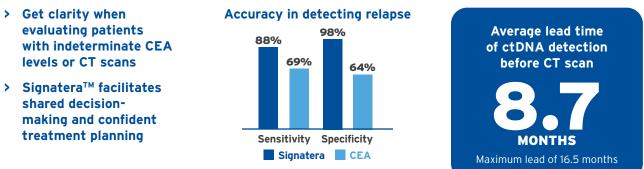
Signatera[™] MRD status outperforms known clinicopathologic risk factors in predicting relapse¹⁻⁴



In the surveillance setting

Use along with CEA testing and other surveillance modalities to detect recurrence earlier, to enable surgical resection or other early intervention

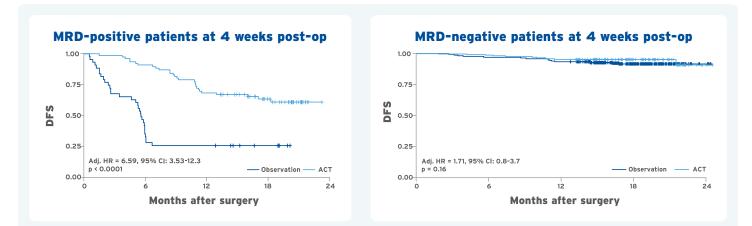
Signatera[™] was shown to detect relapse more accurately than CEA with clinically meaningful lead times over CT scans¹



CEA = carcinoembryonic antigen; CT = computed tomography; ctDNA = circulating-tumour DNA

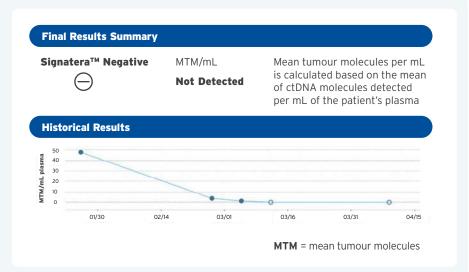
A large-scale prospective, MRD-guided study evaluated the clinical utility of ctDNA analysis in colorectal cancer (CRC)

MRD-positive CRC patients at 4 weeks post-op benefited significantly from chemotherapy while MRD-negative patients at 4 weeks post-op did not demonstrate any significant trend in treatment benefit⁵



Track ctDNA dynamics to enable longitudinal monitoring

> Signatera[™] reports presence/ absence of ctDNA and ctDNA quantity in terms of MTM/mL for longitudinal assessment



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Signatera™ Residual disease test (MRD)



Just like no two tumours are alike, Signatera[™] is personalized for each patient



Tumour-informed MRD assay for individualized care

Customized for each patient's unique tumour signature using Whole Exome Sequencing (WES) to target the top clonal mutations

High sensitivity and specificity for accurate MRD assessment

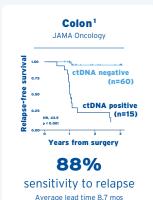
- > By only tracking tumour-specific variants, sensitivity is optimized with a LOD down to 0.01% VAF⁶
- > Filters out germline and CHIP mutations to reduce background noise and to minimize false positives

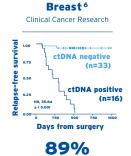
Reliable longitudinal monitoring for confident decision-making

- > Tracks ctDNA dynamics by MTM/mL to enable longitudinal monitoring with a simple blood draw
- Follows clonal mutations that should persist as the tumour evolves

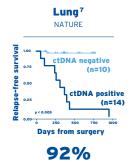
LOD = limit of detection; CHIP = clonal hematopoiesis of indeterminate potential; VAF = Varient allele frequency; WES = whole exome sequencing

Signatera[™] is validated across multiple tumour types^{1,6-8}





sensitivity to relapse Average lead time 9.5 mos

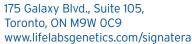


sensitivity to relapse Average lead time 4.0 mos



sensitivity to relapse Average lead time 2.8 mos

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Residual disease test (MRD)

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- 5. Kotani D. et al., Molecular residual disease and efficacy of adjuvant chemotherapy in patients with colorectal cancer, Nature Medicine v29 Issue 1 Jan 2023
- Coombes RC, Page K, Salari R, et al. Personalized Detection of Circulating Tumour DNA Antedates Breast Cancer Metastatic Recurrence. Clin Cancer Res. 2019;25(14):4255-4263.

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contact our Genetics Team

Ask.Genetics@lifelabs.com

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- 8. Christensen E, Birkenkamp-Demtroder K, Sethi H, et al. Early Detection of Metastatic Relapse and Monitoring of Therapeutic Efficacy by Ultra-Deep Sequencing of Plasma Cell-Free DNA in Patients With Urothelial Bladder Carcinoma. J Clin Oncol. 2019;37(18):1547-1557.
- 9. Data on file

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Reduce chemotherapy in colorectal cancer

Signatera™

May help up to **30%** of colorectal cancer **patients** avoid unnecessary **chemotherapy**

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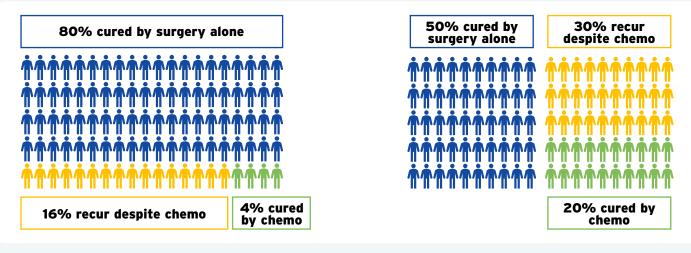
Signatera™ Residual disease test (MRD)

LIFELADS GENETOCS®

Signatera in the Colorectal Cancer (CRC) adjuvant chemotherapy setting

Stage II







Results from clinical trials suggest that assessing for residual disease by Signatera can help avoid unnecessary adjuvant chemotherapy treatment in up to 30% of CRC patients¹ and decrease debilitating toxicities such as peripheral neuropathy²

Signatera is available in Canada:

- Order online (ON residents)
- Order online (BC residents)
- & All Canadians can order over the phone: 1-844-363-4357
- Or via email: <u>Ask.Genetics@LifeLabs.com</u>

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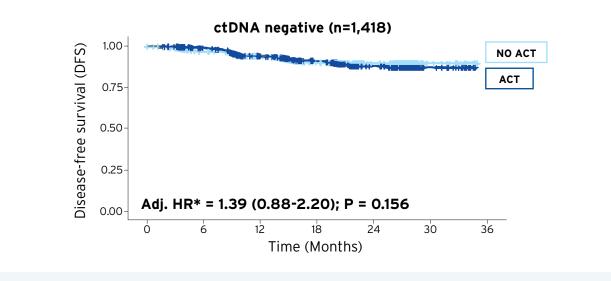


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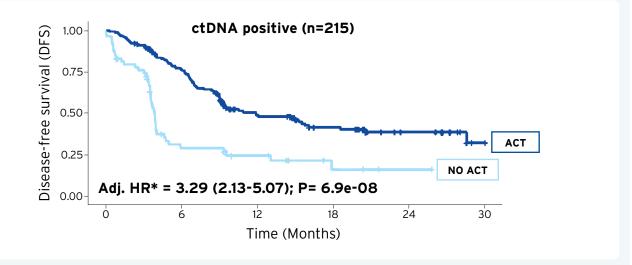


Signatera can inform treatment decisions

Stage II-III CRC patients who **remain ctDNA negative** show **no benefit** from additional **adjuvant chemotherapy (ACT)** 36 months after surgery²



Stage II-III CRC patients who **remain ctDNA positive benefit** from **ACT**, with 37% disease-free survival vs. only 16% in those with no ACT 24 months after surgery²



References

1. Natera, LifeLabs research and analysis

*adjusted for age, gender, MSI, pathological stage, and performance status

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- an updated 24 months (mos) disease free survival (DFS) analysis from GALAXY study (CIRCULATE-Japan). ESMO. 2023.

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