

## Improved liquid biopsy assay performance using sequencing by binding (SBB)

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LIN-28, a recently identified gene, has been found to play a role in the transformation of somatic  
Liquid biopsy is revolutionizing the field of early cancer detection research through non-invasive  
detection of tumor DNA in the blood. However, existing liquid biopsy assays are limited in their  
sensitivity for ctDNA detection at low variant allele frequencies (VAFs). Here, we describe the  
application of the PacBio Onso short-read sequencing system to enable detection of ctDNA at low  
VAFs using the SeraCare Complete ctDNA Mutation Mix reference sample.

We observe superior sensitivity for ctDNA detection using SBB compared to SBS at low VAFs (0.05%,  
0.10%) at comparable sequencing depth. Furthermore, SBB requires significantly less sequencing to  
achieve comparable sensitivity results to SBS. Taken together, our results demonstrate the potential  
of SBB to improve upon existing liquid biopsy methods and better enable research on early cancer  
detection.