

ctDNA analysis of *EGFR*-mutant NSCLC patients receiving osimertinib following previous tyrosine kinase inhibitor treatment

Jamie J Beagan^{1*}, Sander Bach^{2*}, Robert A A van Boerdonk¹, Erik van Dijk¹, Erik Thunnissen¹, Daan van den Broek³, Janneke Weiss⁴, Geert Kazemier², D Michiel Pegtel¹, Idris Bahce⁵, Bauke Ylstra^{1#} and Daniëlle A M Heideman^{1#}

Amsterdam UMC, Vrije Universiteit Amsterdam, Pathology¹, Surgery², Clinical Genetics⁴, Pulmonary Diseases⁵, De Boelelaan 1117, Amsterdam, The Netherlands
Department of Laboratory Medicine³, The Netherlands Cancer Institute, Amsterdam, The Netherlands

*# Contributed equally as first and senior authors, respectively

Aim

Explore the feasibility of ctDNA testing in a clinical setting for NSCLC patients receiving osimertinib as a second or third line EGFR-TKI

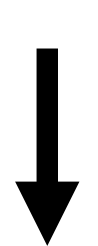
Background

- Circulating tumor (ct)DNA analysis is increasingly used to guide clinical management of advanced non-small cell lung cancer (NSCLC)
- Clinically-actionable *EGFR* mutations can be detected in ctDNA before or after first-line EGFR-Tyrosine Kinase Inhibitor (TKI) treatment
- Data concerning ctDNA detection are limited for patients with a complex treatment history

Materials and Methods

Twenty NSCLC patients who had received osimertinib as a second or third line EGFR-TKI. Pathological and clinical data were acquired.

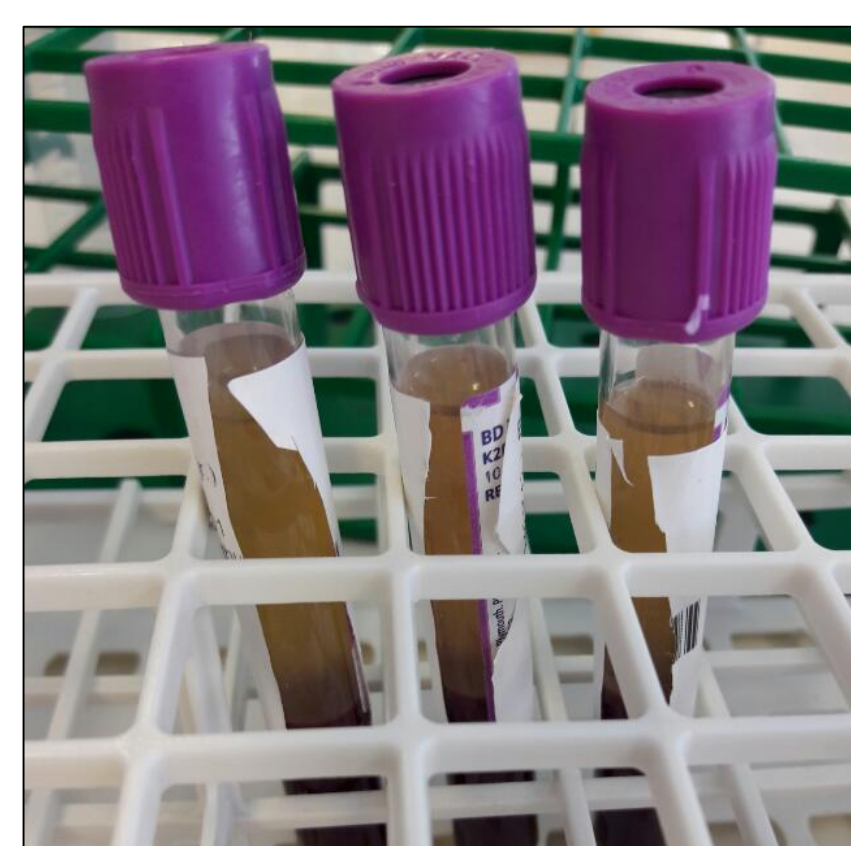
Tumor-tissue



Illumina TruSeq Amplicon Cancer Panel (48 genes)



Plasma cfDNA



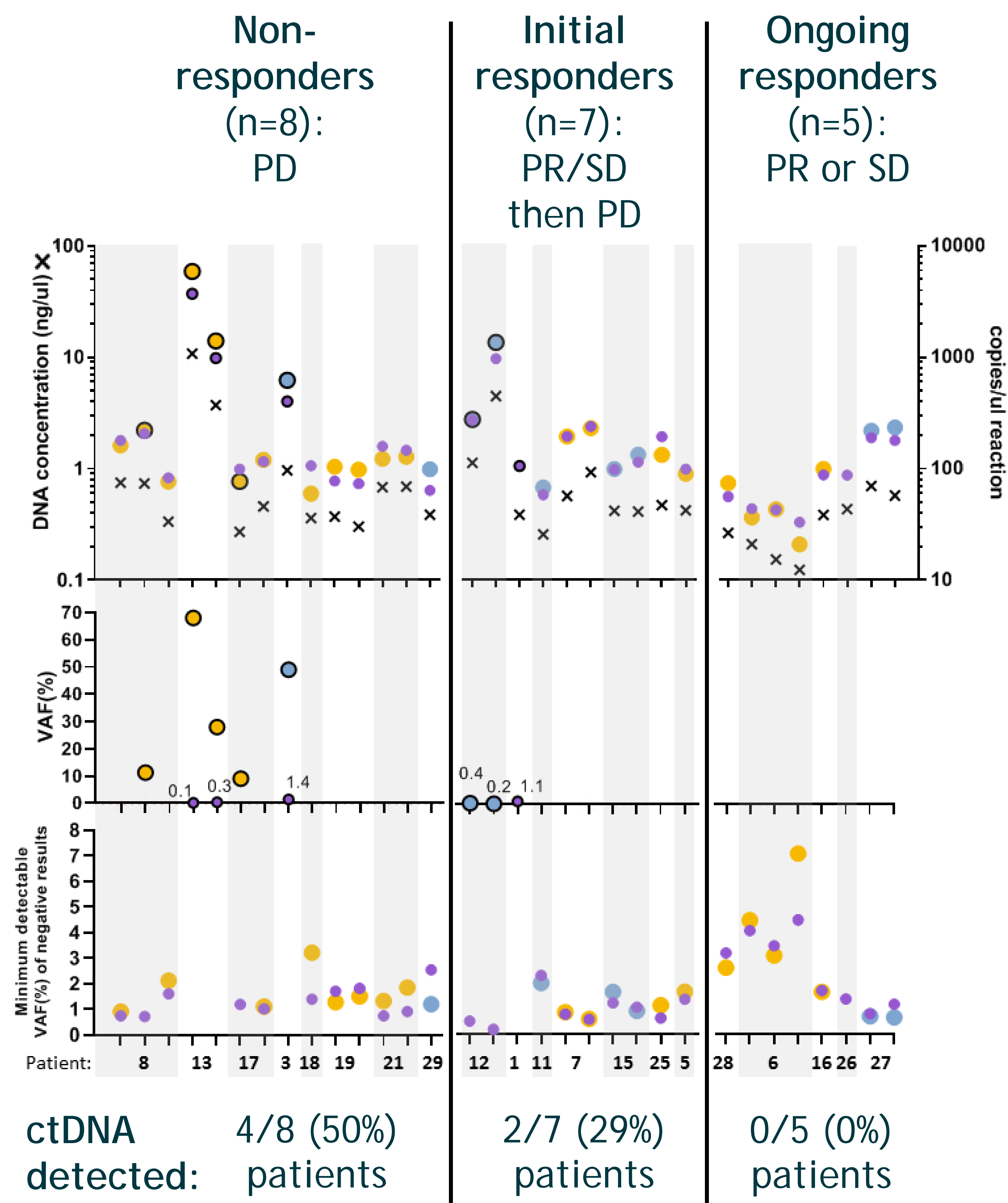
EGFR mutations by Droplet digital PCR (ddPCR)



Results

Patients classified based on osimertinib response

- Key:
- PD: Progressive Disease
 - PR: Partial Response
 - SD: Stable Disease
 - Exon 19 del
 - L858R
 - T790M
- wt mutant



The fraction of *EGFR*-mutant ctDNA tended to be higher in non-responders (0.1 - 68%) than initial responders (0.2 - 1.1%).

No significant difference in overall cfDNA concentration between response groups or the number of metastatic sites.

Conclusion

These results support a potential role for ctDNA analysis in response monitoring of NSCLC patients with a complex EGFR-TKI treatment history.

