

The Assay Matters: ctDNA Assessment in Localized Pancreatic Cancer



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Introduction

- While circulating tumor DNA, (ctDNA) has emerged as a pancreatic cancer (PDAC) biomarker candidate in other malignancies, its investigation is emerging in localized PDAC.
- Both digital droplet PCR (ddPCR) and next generation sequencing (NGS) are published methods of assessing ctDNA
- The advantages and disadvantages of ctDNA assessment by ddPCR and NGS in patients with localized PDAC treated with NAC are unclear

Objectives

- To assess detection and prognostic capabilities of PDAC ctDNA assessment by NGS and ddPCR
- To compare and examine the relative advantages of PDAC ctDNA assessment by NGS and ddPCR

Methods

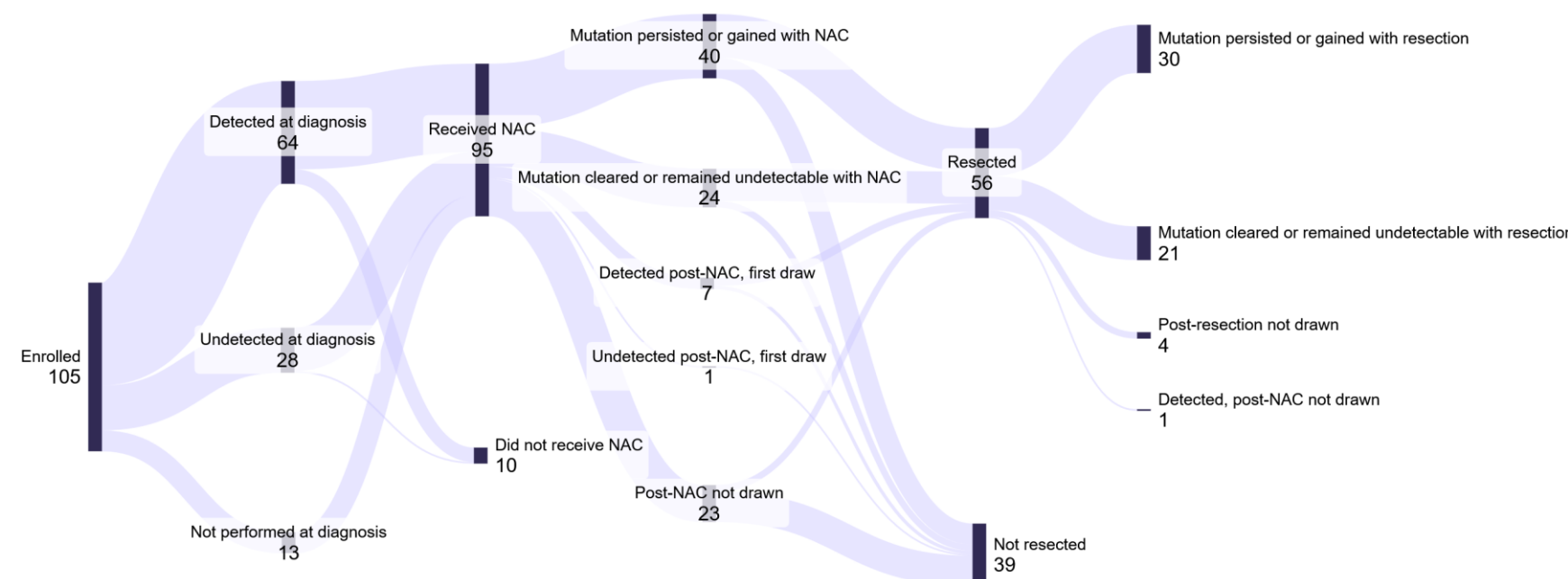
Data source: Prospectively recruited cohort

Inclusion Criteria

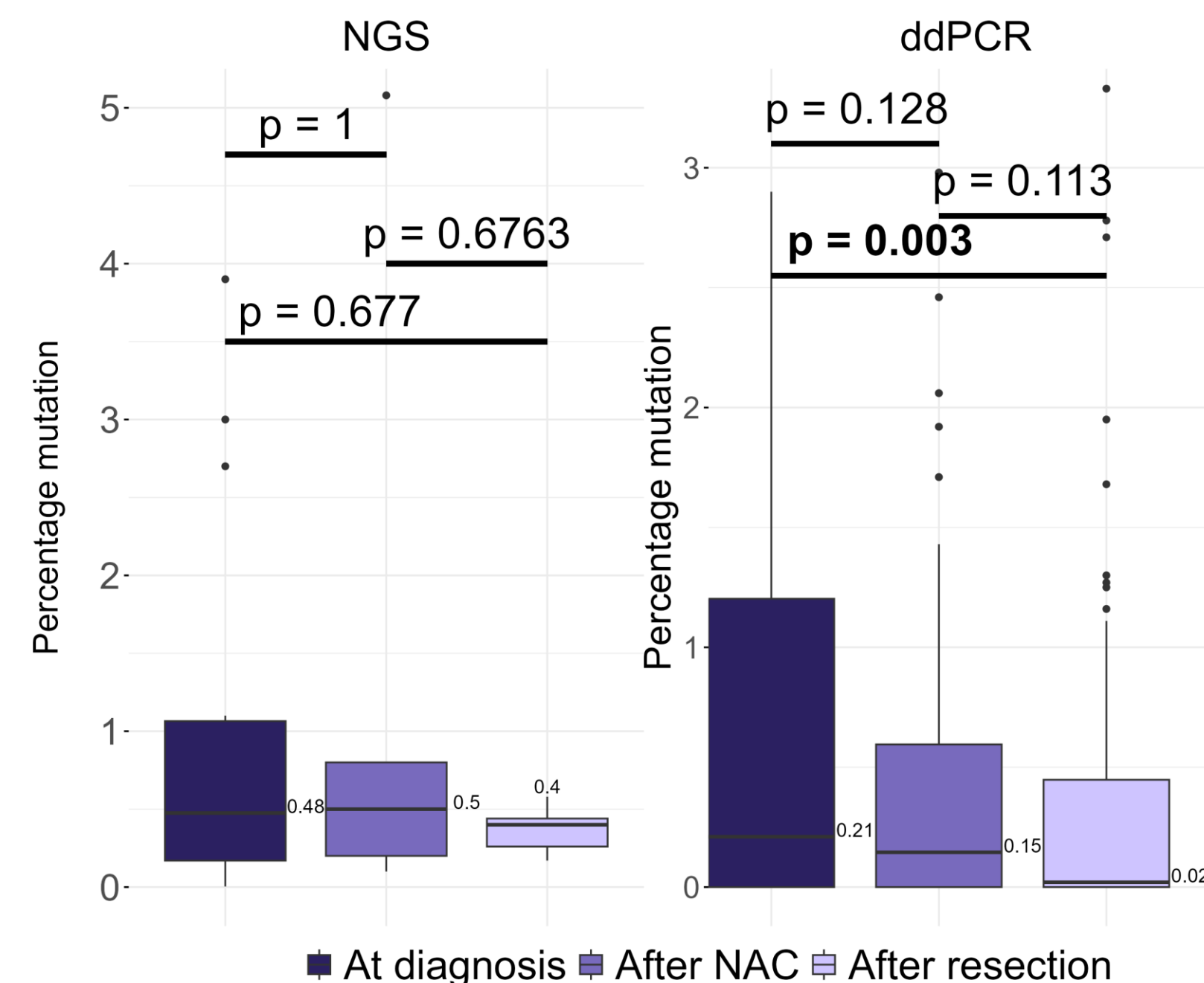
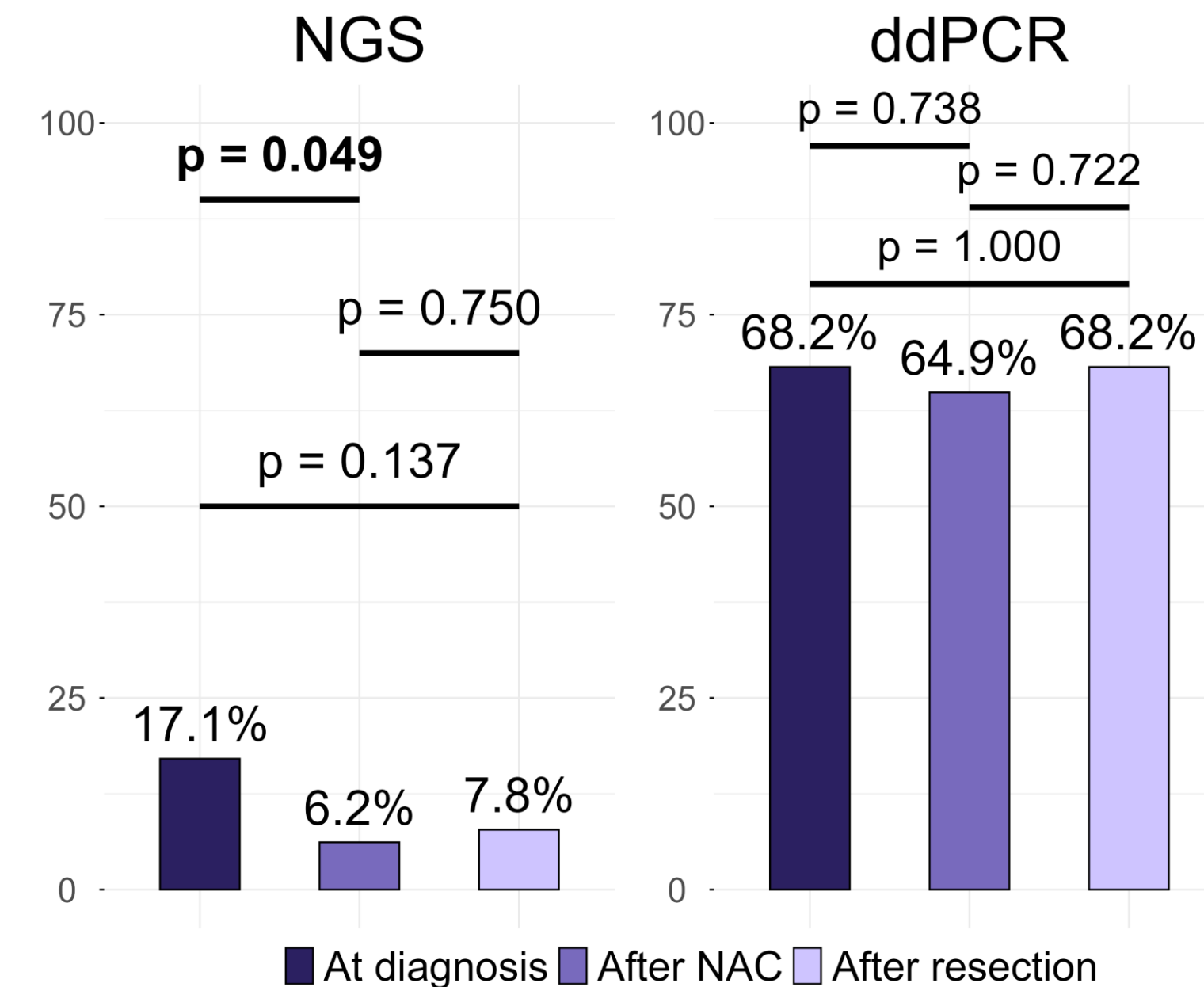
- Newly diagnosed PDAC
- Patients with resectable disease
- Patients planned to undergo NAC

Sample handling

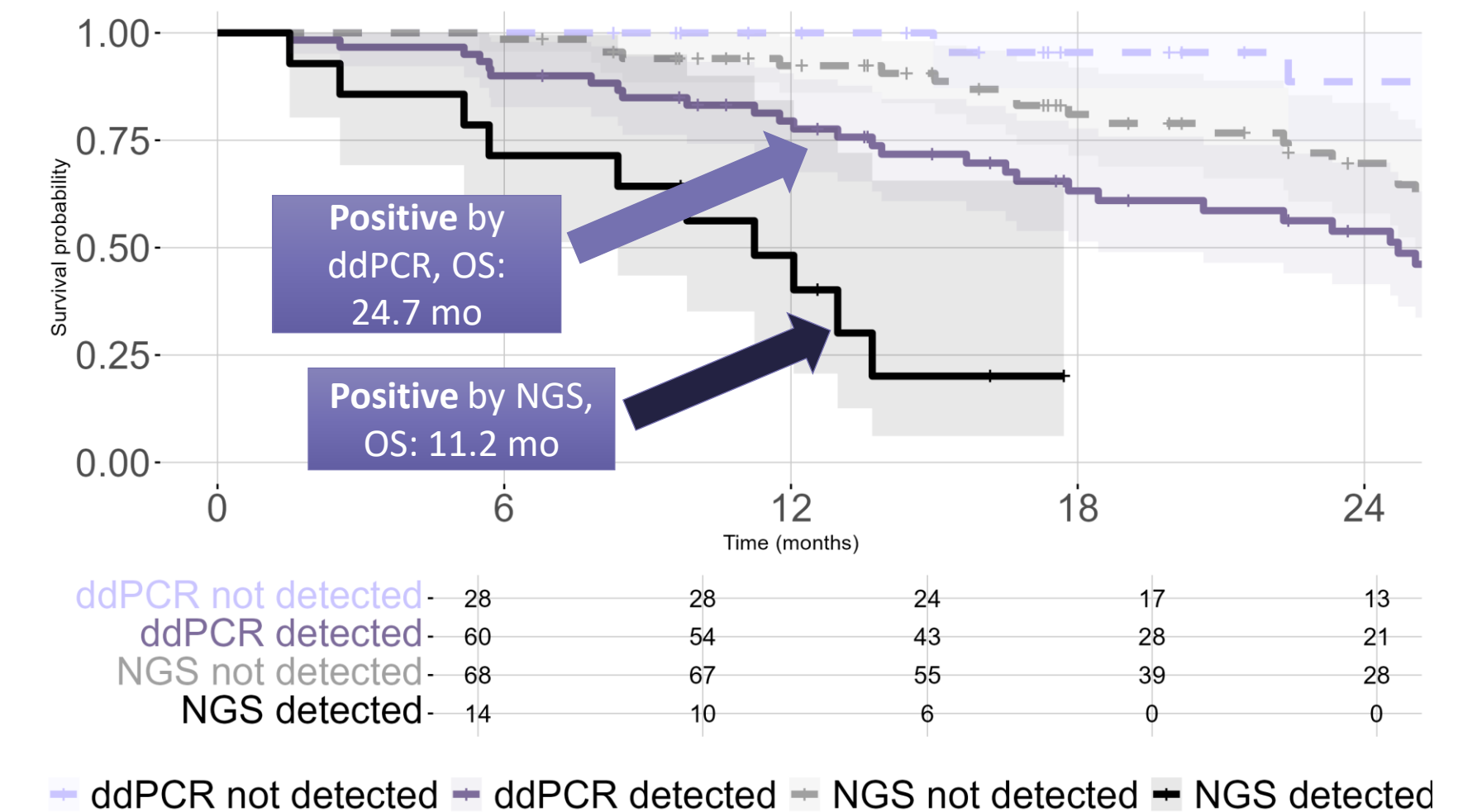
- Peripheral blood samples collected at diagnosis, after NAC, and after resection
- Samples analyzed by ddPCR for mutant *KRAS* G12D, G12V, and G12R



Results – Descriptive Statistics and Regression



Results – Kaplan-Meier Estimates



Conclusions

- Mutant *KRAS* ctDNA is detectable by NGS and ddPCR throughout treatment in patients with localized PDAC
- Detection is prognostic
- ddPCR has higher sensitivity for the detection of smaller ctDNA quantities than NGS
- The higher ctDNA loads required for detection by NGS implicate clinically more aggressive disease

Further study by our group using ddPCR in the detection of mutant *KRAS* ctDNA will include incorporation into adaptive-treatment trial designs such as Alliance A021806

Key points

- The use of ctDNA as a biomarker in PDAC is expanding
- Both NGS and ddPCR were able to detect ctDNA throughout treatment
- ddPCR is more sensitive for ctDNA detection than NGS
- Detection is prognostic
- Future work will include the investigation of ctDNA dynamics during treatment
- Future work will also include investigation of the use of ctDNA to inform NAC regimens and optimally select patients for surgical resection