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Dear Dr. Ridd,

We would like to resubmit our manuscript (NCOMMS-20-25443-T), "The origin, distribution, and genetic interactions of *KRAS* alleles across cancer types," to be considered for publication in *Nature Communications*. We are thankful to the reviewers for their enthusiasm about our study and for their thoughtful comments. We believe we have addressed their comments fully and that the manuscript has improved significantly as a result. In response to the reviewers' comments, our revised manuscript includes:

- 1. A new analysis, presented in Fig 2b.
- 2. Three new supplementary figures.
- 3. Five new supplementary tables.

We confirm that this manuscript has not been published elsewhere and is not under consideration by another journal. All authors have approved the manuscript and agree with its submission to *Nature Communications*. We look forward to your decision.

Sincerely.

Kevin M. Haigis
Associate Professor of Medicine
Dana-Farber Cancer Institute

Harvard Medical School

Peter Park
Professor of Medical Informatics
Harvard Medical School

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