**Title: MR-TRYX platform exploits horizontal pleiotropy to infer novel causal pathways within a Mendelian randomization framework**

We are grateful for the comments that we received in response to the resubmission.

**REVIEWERS' COMMENTS:**  
  
**Reviewer #1 (Remarks to the Author):**  
I think the authors did an excellent job in responding to my comments and in revising the text. I feel the new version of the manuscript is much improved in terms of clarity and scope, and recommend publication.  
  
  
**Reviewer #2 (Remarks to the Author):**  
The authors have adequately addressed my comments.  
  
  
**Reviewer #3 (Remarks to the Author):**  
I believe my concerns have now been adequately addressed. The paper is clearer and I appreciate the added results and comments from the authors which improve the paper in my opinion. I was particularly interested to read the additional results about the LASSO selection procedure.