

# NPDR Supplementary Material

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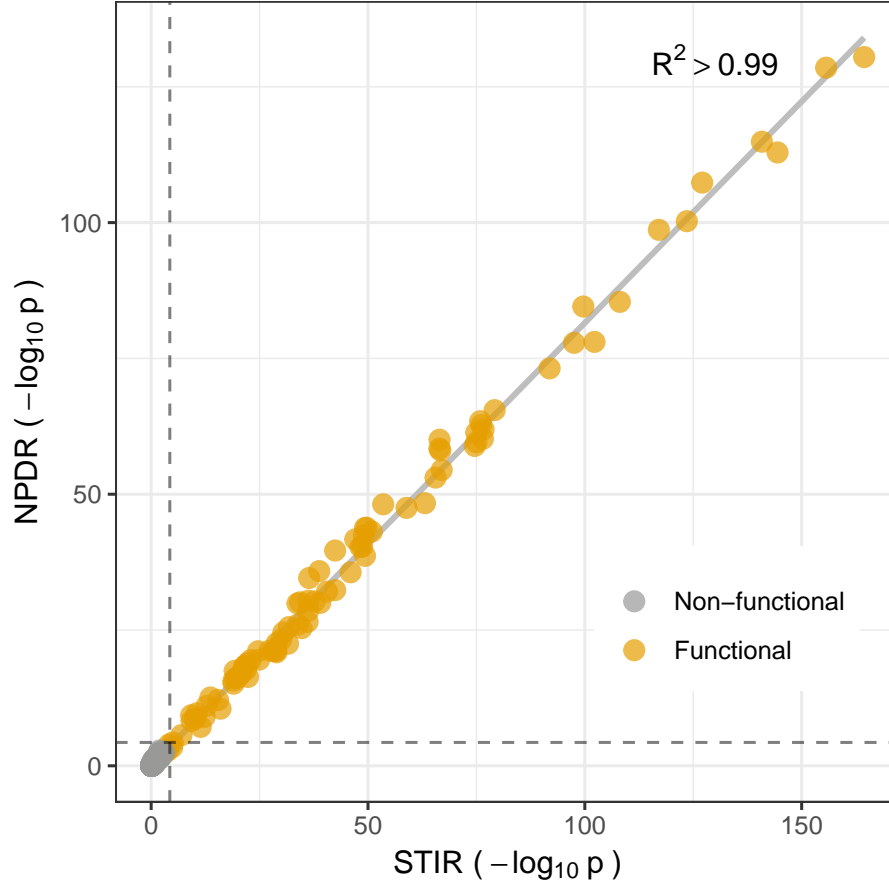


Figure S1: Similarity between NPDR and STIR for one simulation of  $m = 200$  samples and  $p = 1000$  attributes. In 100 replications,  $R^2$  ranges from 0.9827 to 0.9994. STIR is based on a t-test of projected distances and NPDR is based on a logistic regression of projected distances.

## References

- [1] Trang T. Le, Jonathan Savitz, Hideo Suzuki, Masaya Misaki, T. Kent Teague, Bill C. White, Julie H. Marino, Graham Wiley, Patrick M. Gaffney, Wayne C. Drevets, Brett A. McKinney, and Jerzy Bodurka. Identification and replication of RNA-Seq gene network modules associated with depression severity. *Translational Psychiatry*, 8(1):180, September 2018.
- [2] Trang T Le, Ryan J Urbanowicz, Jason H Moore, and Brett A McKinney. Statistical inference relief (stir) feature selection. *Bioinformatics*, 2018.

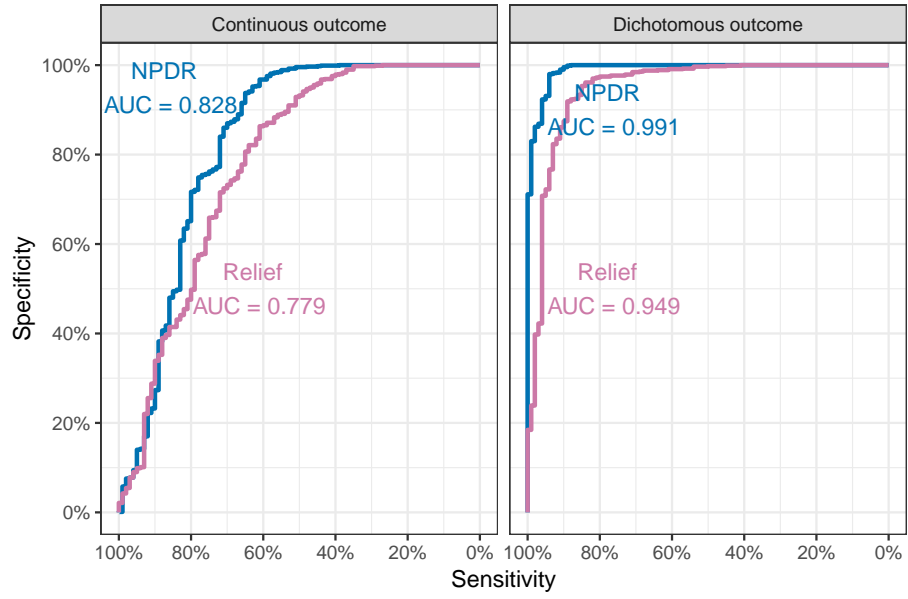


Figure S2: Receiver Operating Characteristics (ROC) curves for Relief-F and NPDR for simulated case-control data with interactions (left) and RRelief and NPDR for simulated continuous outcome data with main effects (right). Simulation uses  $m = 200$  samples and  $p = 1000$  attributes with 100 functional.

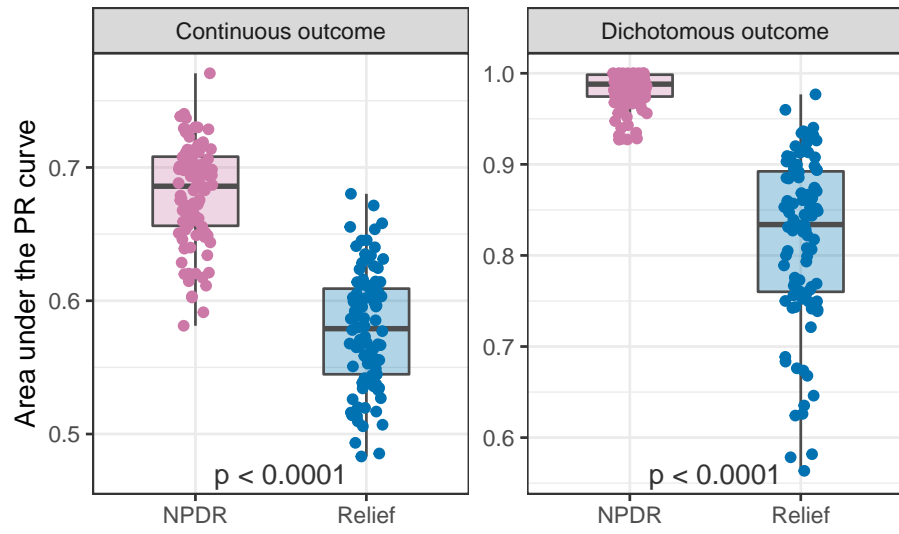


Figure S3: NPDR and Relief comparison of area under the PRC for 100 replicate simulations of case-control (left) and continuous (right) data. All simulations use  $m = 200$  samples and  $p = 1000$  attributes with 100 functional. NPDR yields significantly higher auPRC.

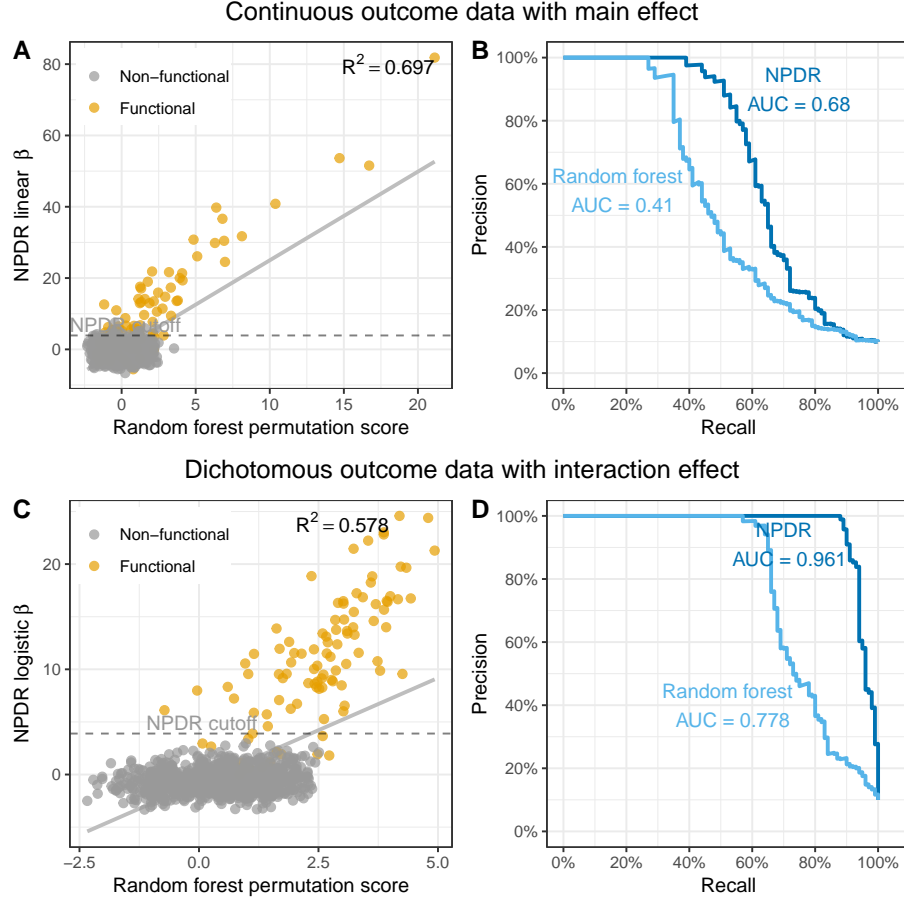


Figure S4: **Comparison of NPDR and random forest importance scores** for continuous outcome data with main effects (top row) and dichotomous outcome data with interaction effects (bottom row). Results for one replicate simulation ( $m = 200$  samples and  $p = 1,000$  attributes with 100 functional). For continuous outcome (A), importance scores computed by random forest permutation (percent increase in MSE) and NPDR standardized linear regression coefficient. For case-control outcome (C), scores computed by random forest permutation (mean decrease in accuracy) and NPDR standardized logistic regression coefficient. A regression line between the scores with  $R^2$  is shown, and a 0.05 Bonferroni cutoff (dashed) is shown for NPDR (A and C). There is no statistical threshold for random forest, so area under the precision-recall curve (auPRC) is used to compare algorithm performance (B, D).

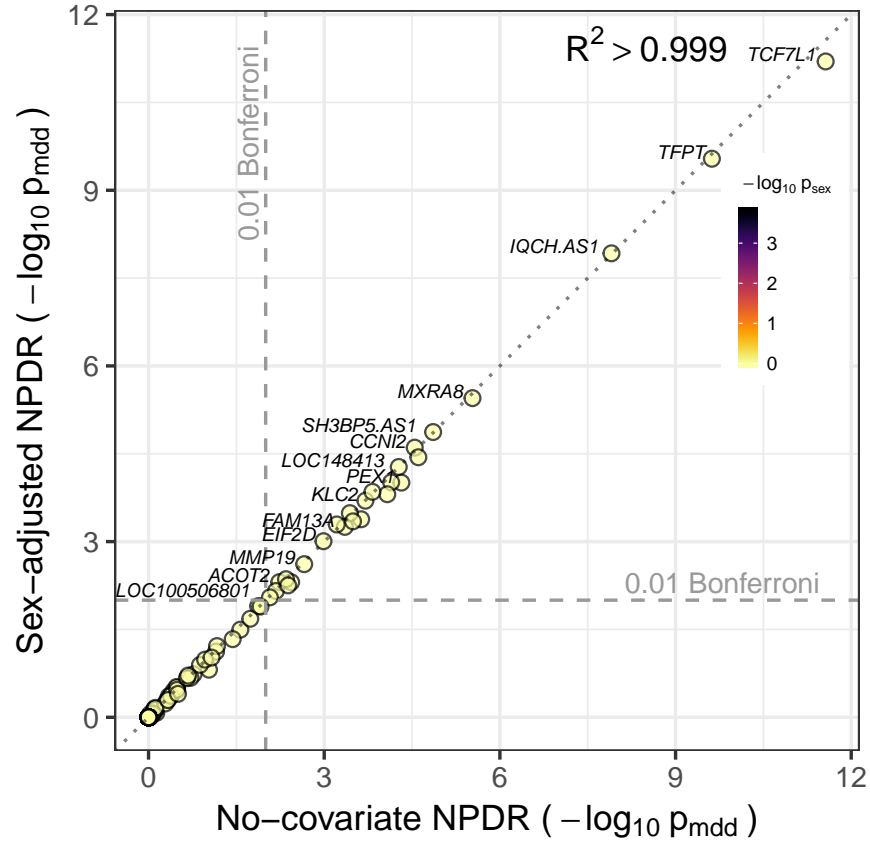


Figure S5: NPDR with and without sex adjustment for analysis of MDD-associated genes in Le et al.'s RNASeq dataset [1]. Adjustment for the sex covariate has a negligible effect on the resulting P values for each important gene because of the balanced study design. Both methods yield consistent results with STIR from previous study (Fig. 4 of Ref. [2]), not shown.

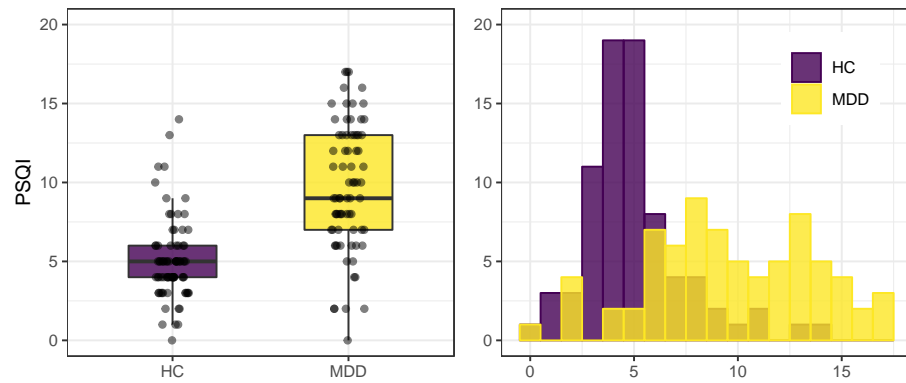


Figure S6: The distribution of the Pittsburgh Sleep Quality Index (PSQI) among individuals with and without MDD.