

- BGmisc: An R Package for Extended Behavior
 Genetics Analysis
- S. Mason Garrison 1, Michael D. Hunter 2, Xuanyu Lyu 1,3,4, Jonathan D. Trattner 1, and S. Alexandra Burt 3
- 1 Wake Forest University, North Carolina, USA 2 Pennsylvania State University, Pennsylvania, USA 3
- 6 Institute for Behavioral Genetics, University of Colorado at Boulder, Boulder, USA 4 Department of
- Psychology & Neuroscience, University of Colorado at Boulder, Boulder, USA 5 Michigan State
- 8 University, Michigan, USA

DOI: 10.xxxxx/draft

Software

- Review 🗗
- Repository [™]
- Archive 🗗

Editor: Open Journals ♂ Reviewers:

@openjournals

Submitted: 01 January 1970 **Published:** unpublished

License

Authors of papers retain copyright and release the work under a ²¹ Creative Commons Attribution 4.0 International License (CC BY 4.0)

Summary

13

Behavior genetics focuses on understanding genetic and environmental influences on individual differences. Traditionally, twin studies have been at the forefront of this discipline. However, research has moved beyond the classical twin design to embrace more complex family structures such as children of twins (CoT) (D'Onofrio et al., 2003), mother-daughter-aunt-niece (MDAN) (Rodgers et al., 2008), and other extended family designs. This expansion allows for a deeper, more nuanced exploration of genetic and environmental influences, but it also introduces challenges, particularly in data structuring and modeling.

A notable challenge arises when researchers venture into extended family data. The data structures inherent in such family designs are orders of magnitude larger than traditional designs. In the classical twin study, a family will consist of a single pair of twins (i.e., two people), whereas in the MADN design a family consists of two mother-daughter pairs (i.e. four people). This problem becomes intractable, when applied to pedigrees, where a single family can be of any size. The BGmisc package addresses this gap by offering a comprehensive suite of functions for structuring and modeling such data.

Statement of need

As behavior genetics delves into more complex data structures like pedigrees, the limitations of current tools become evident. The BGmisc R package addresses these challenges, going beyond what is available in tools like OpenMx and EasyMx that mainly focus on classical twin models.

Two widely used R packages in genetics modeling are OpenMx (Neale et al., 2016) and kinship2 (J. P. Sinnwell et al., 2014; J. Sinnwell & Therneau, 2022). The OpenMx (Neale et al., 2016) package is a workhorse in behavior genetic research. Not only is it a general-purpose software for structural equation modeling that is popular among behavior geneticists (Garrison, 2018), but also for its unique features — the mxCheckIdentification() function. This function checks whether a model is identified, determining if there is a unique solution to estimate the model's parameters based on the observed data. In addition, EasyMx (Hunter, 2023) is a more user-friendly package that streamlines the process of building and estimating structural equation models. It seamlessly integrates with OpenMx's infrastructure. Its functionalities range from foundational matrix builders like emxCholeskyVariance and emxGeneticFactorVariance to more specialized functions like emxTwinModel designed for classical twin models. Despite their strengths, EasyMx and OpenMx have limitations when handling extended family data. Notably, they lack functions for handling modern molecular designs (Kirkpatrick et al., 2021),



- 42 modeling genetic complex relationships, inferring relatedness, or simulating pedigrees.
- Although not a staple in behavior genetics, the kinship2 (J. P. Sinnwell et al., 2014) package provides core features to the broader statistical genetics scientific community, such as plotting pedigrees and computing genetic relatedness matrices. It uses the Lange algorithm (Lange, 2002) to compute relatedness coefficients. This recursive algorithm is discussed in great detail elsewhere, laying out several boundary conditions and recurrence rules. The BGmisc package
- extends the capabilities of kinship2 by introducing an alternative algorithm to calculate the
- relatedness coefficient based on network models. By applying classic path-tracing rules to the entire network, this new method is computationally more efficient by eliminating the need for
- a multi-step recursive approach.

52 Features

57

58

61

62

65

68

70

71

The BGmisc package offers various features tailored for extended behavior genetics analysis.

These features are grouped under two main categories, mirroring the structure presented in our vignettes.

56 Modeling and Relatedness:

- Model Identification: BGmisc evaluates whether a variance components model is identified
 and fits the model's estimated variance components to observed covariance data. The
 technical aspects related to model identification have been described in Hunter et al.
 (2021).
- Relatedness Coefficient Calculation: Using path tracing rules first described in (Wright, 1922) and formalized in (McArdle & McDonald, 1984), BGmisc calculates the (sparse) relatedness coefficients between all pairs of individuals in extended pedigrees based purely on mother and father identifiers.
- Relatedness Inference: BGmisc infers the relatedness between two groups based on their observed total correlation, given additive genetic and shared environmental parameters.

Pedigree Analysis and Simulation:

- Pedigree Conversion: BGmisc converts pedigrees into various relatedness matrices, including additive genetics, mitochondrial, common nuclear, and extended environmental relatedness matrices.
- Pedigree Simulation: BGmisc simulates pedigrees based on parameters including the number of children per mate, generations, sex ratio of newborns, and mating rate.
- Collectively, these tools provide a valuable resource for behavior geneticists and others who work with extended family data. They were developed as part of a grant and have been used in several ongoing projects (Burt, 2023; Garrison et al., 2023; Hunter et al., 2023; Lyu et al., 2023), and theses (Lyu, 2023).

Availability

The BGmisc package is open-source and available on both GitHub at https://github.com/
R-Computing-Lab/BGmisc and the Comprehensive R Archive Network (CRAN) at https:
//cran.r-project.org/package=BGmisc. It is licensed under the GNU General Public License.

Acknowledgments

- $_{82}$ The current research is supported by the National Institute on Aging (NIA), RF1-AG073189.
- We want to acknowledge assistance from Carlos Santos.



References

- Burt, S. A. (2023). Mom genes: Leveraging maternal lineage to estimate the contributions of mitochondrial DNA. *Behavior Genetics*.
- D'Onofrio, B. M., Turkheimer, E. N., Eaves, L. J., Corey, L. A., Berg, K., Solaas, M. H., & Emery, R. E. (2003). The role of the children of twins design in elucidating causal relations between parent characteristics and child outcomes. *Journal of Child Psychology and Psychiatry*, 44(8), 1130–1144. https://doi.org/ff3dbm
- Garrison, S. M. (2018). Popular Structural Equation Modeling Programs for Behavior Genetics.
 Structural Equation Modeling: A Multidisciplinary Journal, 25(6), 972–977. https://doi.org/10.1080/10705511.2018.1493385
- Garrison, S. M., Lyu, X., Hunter, M. D., Rodgers, J. L., Smith, K. R., Coon, H., & Burt, S.
 A. (2023). Analyzing extended cousin similarity to unravel the mystery of mtDNA and longevity. *Behavior Genetics*.
- Hunter, M. D. (2023). EasyMx: Easy model-builder functions for 'OpenMx'. https://CRAN.
 R-project.org/package=EasyMx
- Hunter, M. D., Garrison, S. M., Burt, S. A., & Rodgers, J. L. (2021). The Analytic Identification of Variance Component Models Common to Behavior Genetics. *Behavior Genetics*, 51(4), 425–437. https://doi.org/10.1007/s10519-021-10055-x
- Hunter, M. D., Lyu, X., Garrison, S. M., Rodgers, J. L., Smith, K., Coon, H., & Burt, S. A. (2023). Modeling mtDNA effects from extended pedigrees in the utah population database.

 Behavior Genetics.
- Kirkpatrick, R. M., Pritikin, J. N., Hunter, M. D., & Neale, M. C. (2021). Combining Structural-Equation Modeling with Genomic-Relatedness-Matrix Restricted Maximum Likelihood in OpenMx. Behavior Genetics, 51(3), 331–342. https://doi.org/10.1007/s10519-020-10037-5
- Lange, K. (2002). Genetic Identity Coefficients. In K. Lange (Ed.), *Mathematical and Statistical Methods for Genetic Analysis* (pp. 81–96). Springer. https://doi.org/10.1007/978-0-387-21750-5_5
- Lyu, X. (2023). Statistical power analysis on mtDNA effects estimation [Master's thesis].
 Wake Forest University.
- Lyu, X., Hunter, M. D., Rodgers, J. L., Smith, K. R., Coon, H., Burt, S. A., & Garrison, S. M. (2023). Statistical power analysis on mtDNA effects estimation. *Behavior Genetics*.
- McArdle, J. J., & McDonald, R. P. (1984). Some algebraic properties of the reticular action
 model for moment structures. *British Journal of Mathematical and Statistical Psychology*,
 37, 234–251. https://doi.org/10.1111/j.2044-8317.1984.tb00802.x
- Neale, M. C., Hunter, M. D., Pritikin, J. N., Zahery, M., Brick, T. R., Kirkpatrick, R. M., Estabrook, R., Bates, T. C., Maes, H. H., & Boker, S. M. (2016). OpenMx 2.0: Extended Structural Equation and Statistical Modeling. *Psychometrika*, 81(2), 535–549. https://doi.org/f8rfrg
- Rodgers, J. L., Bard, D. E., Johnson, A., D'Onofrio, B., & Miller, W. B. (2008). The Cross-Generational Mother–Daughter–Aunt–Niece Design: Establishing Validity of the MDAN Design with NLSY Fertility Variables. *Behavior Genetics*, 38(6), 567–578. https://doi.org/10.1007/s10519-008-9225-0
- Sinnwell, J. P., Therneau, T. M., & Schaid, D. J. (2014). The kinship2 r package for pedigree data. *Human Heredity*, 78, 91–93. https://doi.org/10.1159/000363105



Sinnwell, J., & Therneau, T. (2022). *kinship2: Pedigree functions*. https://CRAN.R-project.org/package=kinship2

Wright, S. (1922). Coefficients of inbreeding and relationship. The American Naturalist, 56(645), 330–338. https://doi.org/10.1086/279872

