



To Merge Randomized Controlled Trials and Real-world Evidence with Bayesian Network Meta-regression: A Case Study in Patients with Myelodysplastic Syndromes

Li Jiu¹, Junfeng Wang¹, Rick A Vreman^{1,2}, Aukje K Mantel-Teeuwisse¹, Wim G Goettsch^{1,2}

¹ Division of Pharmacoepidemiology and Clinical Pharmacology, Utrecht Institute for Pharmaceutical Sciences, Utrecht University, Utrecht, The Netherlands
² National Health Care Institute, Diemen, The Netherlands

Background & Objective

- Randomized controlled trials (RCTs) and real-world evidence (RWE) are often synthesized separately in health technology assessment (HTA).
- One reason is that RCTs and RWE show great heterogeneity in methodology and risk of bias which makes merging the two data sources technically difficult.
- To address this problem, Bayesian Network Meta-regression (BNMR) models have been applied for evidence synthesis in the HTA setting.
- Hence, we aimed to estimate and compare the performance of existing BNMR models in a case study of Myelodysplastic Syndromes (MDS).

Method - Meta-analysis

Case

- Data source:** Song et al. (2021);
- Target population:** Patients with acute myeloid leukemia and myelodysplastic syndromes;
- Intervention:** Reduced intensity conditioning (RIC);
- Comparator:** Myeloablative conditioning (MAC);
- Outcomes of interest:** Overall survival (Binary outcome).

Identification of BNMR models using the snowballing approach, according to Wohlin (2016)

- Website that supports the snowballing approach;
- Connected Papers;
- Starting from two identified reviews of appraisal tools: Jenkins et al. (2021) & Zhang et al. (2019)
- Eligibility criteria: (1) Bayesian model; (2) The model supported binary outcomes; (3) Codes for running a model were available.

Data collection & preparation

- Characteristics of BNMR models;
- Codes used to run the BNMR models;
- Covariates: age, duration of follow up.

Model running

- R package: Crossnma & R2jags;
- Initial value set in the BNMR models: Null;
- Number of iterations of Bayesian meta-regression: 50000;
- Number of burn-in iterations: 20000;
- Number of Markov chains: 4
- Number of thinning of Markov chains: 1.

Model comparison

- Comparison of mean and confidence interval in a forest plot.

RESULTS

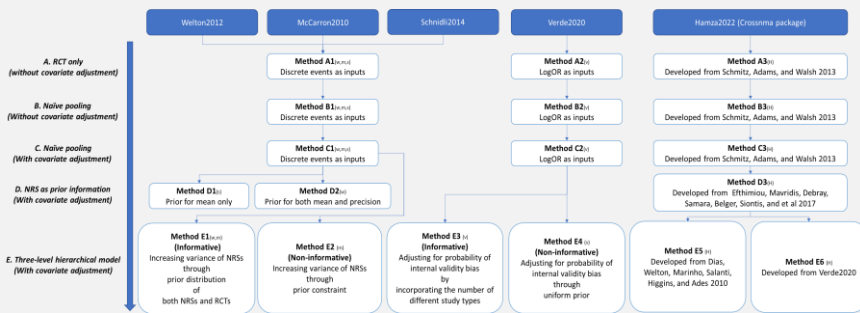


Figure 1. Flow chart of the BNMR models for comparison

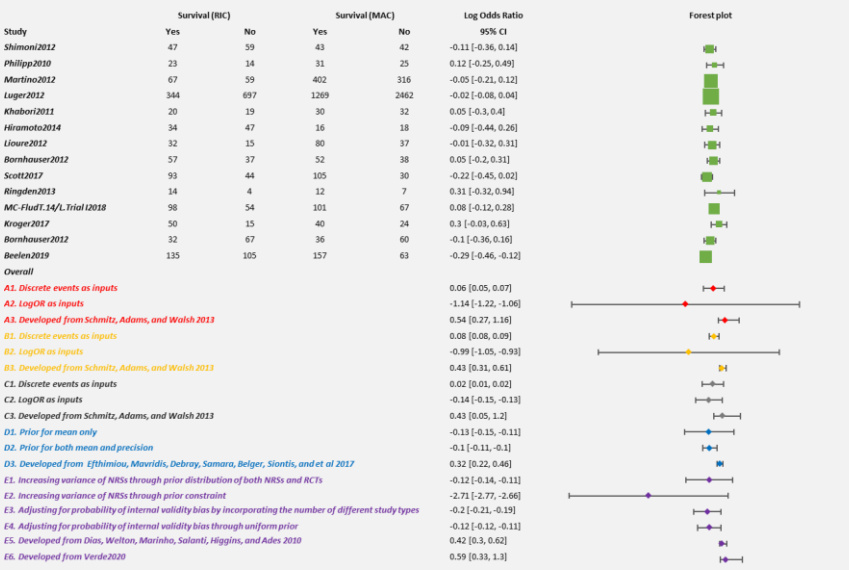


Figure 2. Forest plot

CONCLUSION

- Estimates obtained from the BNMR models are sensitive to model algorithms.
- Further research is needed to confirm our findings by validating these algorithms in other case studies.

Reference

- Song Y, Yin Z, Ding J, Wu T. Reduced Intensity Conditioning Followed by Allogeneic Hematopoietic Stem Cell Transplantation Is a Good Choice for Acute Myeloid Leukemia and Myelodysplastic Syndrome: A Meta-Analysis of Randomized Controlled Trials. *Frontiers in oncology*. 2021 Oct 7;11:708727.
- Jenkins DA, Hussein H, Martina R, Dequen-O'Byrne P, Abrams KR, Bukjiewicz S. Methods for the inclusion of real-world evidence in network meta-analysis. *BMC medical research methodology*. 2021 Dec;21(1):1-9.
- Zhang K, Arora P, Sati N, Bévilleau A, Troke N, Veroniki AA, Rodrigues M, Rios P, Zarin W, Tricco AC. Characteristics and methods of incorporating randomized and nonrandomized evidence in network meta-analyses: a scoping review. *Journal of Clinical Epidemiology*. 2019 Sep 1;113:1-0.
- Wohlin C. Second-generation systematic literature studies using snowballing. *In Proceedings of the 20th International Conference on Evaluation and Assessment in Software Engineering 2016 Jun 1 (pp. 1-6)*.