

INTRODUCTION

- **CohortMethod** performs comparative cohort studies in the Observational Medical Outcomes Partnership (OMOP) common data model. Capabilities include:
 - Propensity models
 - Logistic, Poisson, Cox regression
- We extend **CohortMethod** to include the Fine and Gray model, which extends the Cox proportional hazards model with subdistribution hazard function:

$$\lambda_k(t) = \lim_{\Delta t \rightarrow 0} \frac{\text{Prob}(t \leq T < t + \Delta t, D = k | T \geq t \cup (T < t \cap K \neq k))}{\Delta t}$$

where λ_k is the instantaneous rate of experience the event of interest $D=k$ for subjects, given they have survival to time t without experiencing any competing events $K \neq k$ for observed failure time T .

METHODS

- Kawaguchi, Shen, Li, and Suchard (2019) use a novel forward-backward scan algorithm to linearize computations to reduce complexity from $O(n^2)$ to $O(n)$:
 - Log-pseudo likelihood
 - Gradient
 - Hessian diagonal
- We develop function **combineCompetingStudyPopulations** that combines two study populations, with the outcome of interest and competing event to generate a population with information on subjects experience either outcome
- We include option **riskId** in the function **createFitOutcomeModelArgs** where we specify the competing risk outcome concept ID to fit Fine Gray in the multiple analysis framework
- Regression modelling is fitted using **Cyclops**

We are able to extend **CohortMethod** to perform comparative cohort studies in an observational database using Fine and Gray regression models for competing risks with fast and scalable computational methods.



Take a picture to view the GitHub repository for **OHDSI/CohortMethod@finegray**

RESULTS

- We apply **CohortMethod** to study the relative risk of hospitalization with heart failure for new users under angiotensin-converting enzyme (ACE) inhibitors and thiazide diuretics (THZs).

```
studyPopOutcome <- createStudyPopulation(cohortMethodData = cmData,
                                         outcomeId = 7368,
                                         firstExposureOnly = FALSE,
                                         riskWindowStart = 0,
                                         riskWindowEnd = 9999)

studyPopRisk <- createStudyPopulation(cohortMethodData = cmData,
                                      outcomeId = 7364,
                                      firstExposureOnly = FALSE,
                                      riskWindowStart = 0,
                                      riskWindowEnd = 9999)

studyPopCombined <- combineCompetingStudyPopulations( ## New function
  mainPopulation = studyPopOutcome,
  competingRiskPopulation = studyPopRisk,
  removeSubjectsWithSimultaneousEvents = TRUE
)

ps <- createPs(cohortMethodData = cmData,
              population = studyPopCombined)

matchedPop <- matchOnPs(population = ps,
                       caliper = 0.2)
```

Figure 1. Generating combined study population for both outcomes and performing one-to-one matching on propensity scores.

```
outcome <- fitOutcomeModel(population = matchedPop, ## Updated for multi-type events
                          modelType = "fgr")
outcome

## Model type: fgr
## Stratified: FALSE
## Use covariates: FALSE
## Use inverse probability of treatment weighting: FALSE
## Status: OK
##
## Estimate lower .95 upper .95 logRr seLogRr
## treatment 0.942060 0.873580 1.015845 -0.059686 0.0385
```

Figure 2. Fitting Fine and Gray model on our one-to-one matched population.

REFERENCES

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- Fine JP, Gray RJ. A Proportional Hazards Model for the Subdistribution of a Competing Risk. *Journal of the American Statistical Association*. 1999;94(446):496–509.
- Kawaguchi ES, Shen JI, Li G, Suchard MA. A Fast and Scalable Implementation Method for Competing Risks Data with the R Package fastcmprsk. *The R Journal*. 2020;12(2):163.
- Suchard MA, Simpson SE, Zorych I, Ryan P, Madigan D. Massive Parallelization of Serial Inference Algorithms for a Complex Generalized Linear Model. *ACM Transactions on Modeling and Computer Simulation*. 2013;23(1):1–17