EVALUATION OF THE PARETO FRONTIER APPROACH FOR MODEL CALIBRATION

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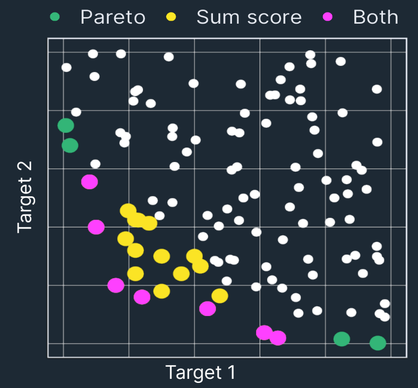
OBJECTIVE: We conducted a simulation study to assess the performance of the Pareto Frontier approach against a conventional distance-based (unweighted) sum score.

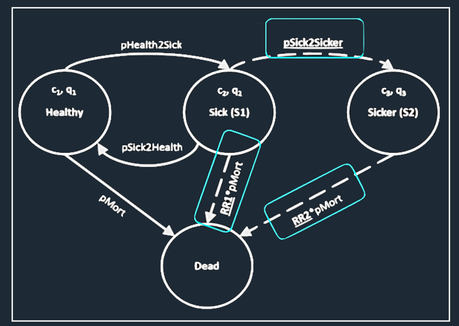
Key reference: Enns EA, Cipriano LE, Simons CT, Kong CY. Identifying best-fitting inputs in health-economic model calibration: a Pareto frontier approach. MDM. 2015 Feb;35(2):170-82.

PARETO FRONTIER

- The Pareto Frontier is model a calibration method, recently proposed by Enns et al. (2015)
- A set of input parameters is on the Frontier, if you cannot improve the fit on one target without reducing it on another (see right figure)



SICK-SICKER MODEL



- We used the same cohort state transition model that Enns et al. presented in their paper (see left figure)
- It has 3 unknown parameters that need to be calibrated
- We tested 4 target sets, consisting of 2-5 targets

IMPLEMENTATION

The study was conducted in R v4.0. The rPref package was used to identify Pareto optimal sets. We used a 64-cores AWS instance and parallelisation to execute the >500 mio. model runs.

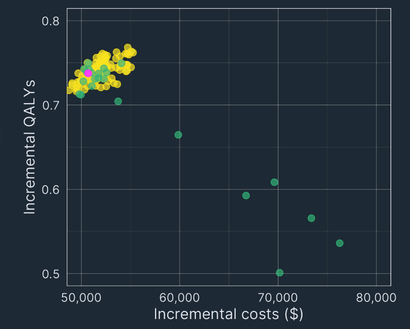
The source code is available at: github.com/bitowaqr/pareto_frontier

SIMULATION PSEUDO CODE

```
for i = 1 to 10,000  {

  1. Specify a true model:
     – Randomly draw values for all (known and unknown) model parameters
     – Compute the true incremental net monetary benefit (iNMB)

  2. Generate calibration target sets:
     – Run a micro-simulation to generate stochastic targets

  3. Run model calibration:
     – Generate 50,000 candidate input sets
     – For each set, compute differences between model outputs and targets
     – For each of the 4 target sets, select the inputs that:
        – lie on the Pareto Frontier
        – are among 1% with the lowest sum of absolute errors

  4. Evaluate calibration performance:
     – Compute the mean iNMB across selected input sets and compare it against the true iNMB

}
```

SAMPLE RESULTS FOR $i = 1$

- The right figure shows exemplary results for one simulation run
- Here, the sum score calibration performed better than the Pareto Frontier approach: the mean absolute error in iNMB was 977 vs. 19,091.



MAIN RESULTS

Mean (SD) absolute error in iNMB

Target set	Pareto Frontier	Sum score
1	11,600 (7,032)	**7,505 (6,381)**
2	7,403 (4,151)	**5,368 (4,104)**
3	**21,095 (8,097)**	28,185 (14,337)
4	12,461 (5,444)	**3,905 (2,859)**

- The sum score method provided more accurate mean iNMB predictions for 3 of 4 target sets
- Models calibrated with the Pareto Frontier approach performed better only when using Target Set 3*
- The mean (SD) number of sets on the Frontier was 601 (984)
- Identifying Pareto optimal inputs was computationally demanding

***Note:** Target set 3 consisted of 3 proportions (range: 0-1) and 1 ratio (range: 0-Inf.). When target trade-offs are (mis-)specified like this, it is not surprising that a sum score performs poorly.

CONCLUSION

1) The Pareto Frontier model calibration method generally performed worse than the simple, distance-based sum score.
2) However, when trade-offs between targets are misspecified, the Pareto Frontier may provide less biased results.

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