

Supplementary material

Supplementary Table 1: Limits for numerical base risk predictors. Values exceeding the lower/upper limit were set to missing. If lower/upper limit is not reported, all values on the lower/upper end were left unchanged.

| | Lower limit | Upper limit | Unit |
|---------------------------------|-------------|-------------|------------|
| Diastolic blood pressure | 40 | | mmHg |
| Haemoglobin | 4 | 20 | g/dL |
| Hematocrit | 15 | | % |
| Leukocytes | | 50000 | 1/ μ L |
| Thrombocytes | 42000 | | 1/ μ L |
| Creatinine | 0.2 | 8.8 | mg/dL |
| Sodium | | 165 | mmol/L |
| Potassium | 2 | 7.5 | mmol/L |
| NT-proBNP | | 35000 | pg/mL |

Supplementary Table 2: Limits for numerical dynamic risk predictors. Values exceeding the lower/upper limit were set to missing. If lower/upper limit is not reported, all values on the lower/upper end were left unchanged.
 * Set to missing in case of atrial fibrillation. ** Set to missing if < 0.9x patient's 5-percentile weight or if > 1.1x patient's 95-percentile weight.

| | Lower limit | Upper limit | Unit |
|------------------------------|-------------|-------------|-------|
| Heart rate during ECG | | | |
| Minimum | 25 | 160 | 1/min |
| Maximum | 40 | 220 | 1/min |
| Average | 40 | 129 | 1/min |
| SpO2 | 75 | | % |
| PQ Interval* | 50 | 675 | ms |
| QRS Interval | 40 | 290 | ms |
| QT Interval | 100 | 780 | ms |
| QT Interval (Bazett) | 100 | 780 | ms |
| Blood pressure | | | |
| Diastolic | 40 | | mmHg |
| Systolic | 70 | | mmHg |
| Average | 50 | | mmHg |
| Weight** | 40 | | kg |

Supplementary Table 3: Heuristic prioritisation rules. Patients are first sorted w.r.t. priority, then w.r.t. points.

| Parameter | Rule | Priority | Points |
|-------------------------|--|----------|--------|
| Weight | 1-day difference > 1 kg | high | 1000 |
| | 0.5 kg <= 1-day difference <= 1 kg | medium | 501 |
| | 2-day difference > 2 kg | high | 1000 |
| | 1.5kg <= 2-day difference <= 2 kg | normal | 501 |
| | 8-day difference > 2.5 kg | critical | 5000 |
| | 2 kg <= 8-day difference <= 2.5 kg | medium | 501 |
| Self-rated well-being | 5 (very bad) | critical | 5000 |
| | 4 (bad) | high | 1000 |
| | 3 (normal) | medium | 501 |
| | Drop from 1 (very good) to 3 (normal) | medium | 501 |
| Blood pressure | Systolic BP < 80 mmHg or Systolic BP > 160 mmHg or Diastolic BP > 100 mmHg | critical | 5000 |
| | Systolic BP < 90 mmHg or Systolic BP > 140 mmHg or Diastolic BP > 90 mmHg or Diastolic BP < 40 mmHg | high | 1000 |
| | 130 mmHg < Systolic BP <= 140 mmHg | medium | 501 |
| Ventricular tachycardia | Any occurrence | critical | 20000 |
| Atrial fibrillation | First occurrence | critical | 10000 |
| | Any other occurrence | medium | 501 |
| Sinus rhythm | Any occurrence | low | 250 |
| Pacemaker rhythm | Any occurrence | low | 250 |
| QRS Interval | First occurrence of QRS Interval < 120 ms (without pacemaker) | critical | 5000 |
| | Any other occurrence of QRS Interval < 120 ms (without pacemaker) | low | 250 |

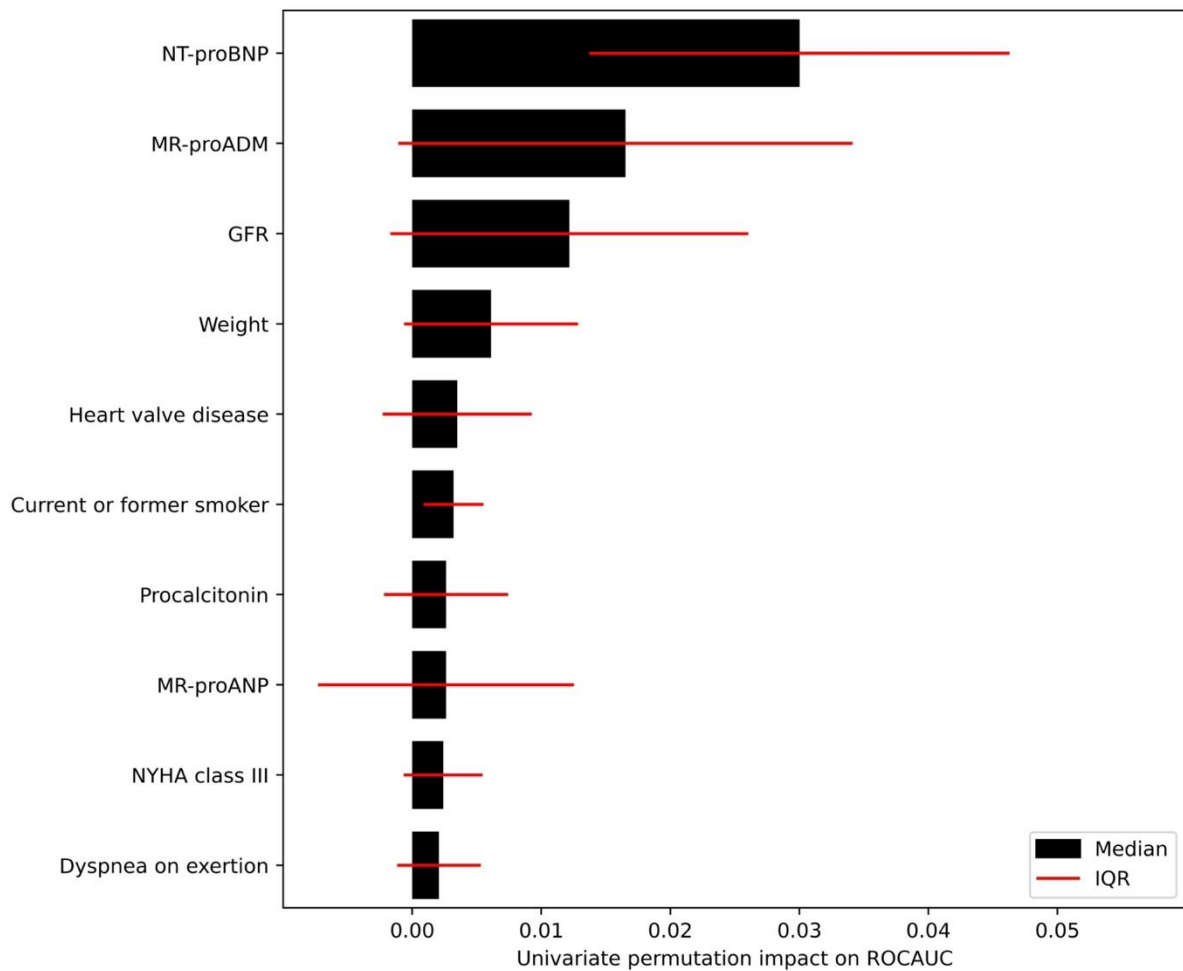
| Parameter | Rule | Priority | Points |
|----------------------|---|----------|--------|
| AV block | None | low | 250 |
| | I | normal | 501 |
| | II | high | 1000 |
| | III | critical | 5000 |
| Heart rate | Minimal heart rate during ECG < 50 beats per minute or > 100 beats per minute | critical | 5000 |
| | Absolute 1-day difference in mean heart rate during ECG > 20 beats per minute | high | 1000 |
| QT Interval (Bazett) | >= 460 ms (without pacemaker) | high | 1000 |
| | >= 480 ms (without pacemaker) | critical | 5000 |
| | 1-day relative difference > 10 % | high | 1000 |
| SpO2 | < 94 % | high | 1000 |
| | < 90 % | critical | 5000 |

Supplementary Table 4: Complete list of predictors included in the final models for the baseline risk and the dynamic risk.

| | Predictors |
|----------------------|-------------------------------------|
| Baseline risk | NT-proBNP |
| | MR-proADM |
| | GFR |
| | Weight |
| | Heart valve disease |
| | Current or former smoker |
| | Procalcitonin |
| | MR-proANP |
| | NYHA class III |
| | Dyspnea on exertion |
| | COPD |
| | Leukocytes |
| | Dilated cardiomyopathy |
| | Age |
| | Hematocrit |
| | HF hospitalisation <= 30 days prior |
| | Haemoglobin |
| | Peripheral artery disease |
| | Diastolic BP |
| | Living in rural area |
| | Body-mass index |
| | Treated with antiplatelet therapy |
| | Sodium |
| | Treated with β -blockers |
| | Stroke |
| | Fluid intake <= 1.5 L/day |

| | |
|---|-------------------------------------|
| | Predictors |
| | LVEF |
| Unplanned HF hospitalisation within 7 days | Base risk score |
| | Self-rated well-being |
| | Weight difference, 13 days |
| | SpO2 |
| | HF hospitalisation <= 30 days prior |
| | Atrial fibrillation |
| | Weight |
| | QT Interval (Bazett's formula) |
| | Systolic BP |
| | PQ Interval |
| | Average heart rate during ECG |
| | Minimal heart rate during ECG |
| | AV block I |
| | Maximal heart rate during ECG |

Supplementary Figure 1: Impact of the 10 highest ranked predictors of the baseline risk. Impact measured by repeatedly permuting each predictor before making predictions and measuring the resulting drop-off in ROCAUC.



S1: Random Forest implementation details

We train a Random Forest of 250 trees without a pre-specified maximum tree-depth. Tuneable hyperparameters are the maximum number of features per tree (between 3 and 8), the minimum number of samples required in a tree-leaf (between 2 and 15), the minimum number of samples required to allow a further tree-split (between 5 and 30), and the fraction of samples used per tree (evenly spaced between 0.7 and 1).

We use 10-fold cross-validation and 50 random draws from the hyperparameter space in the tuning process.

With the optimal choice of hyperparameters, we compute the permutation importance of all predictors. In cross-validation fashion (10 folds), a model is fit on the training folds using all features, and predictions are made on the test fold with each predictor in turn being randomly permuted. Per fold, we repeat the random permutation of each predictor 25 times. The resulting importance score is the median dropoff in ROCAUC (compared with the ROCAUC without permutation) across all folds and permutation-repeats.

We use the permutation importance scores for backwards predictor elimination. Predictors are sorted by their importance score. In ten evenly spaced steps, we eliminate the lowest ranked predictors until only 10 % of all predictors remain. With all ten tested combinations of features, we run hyperparameter tuning, and select the combination of predictors yielding the largest cross-validated ROCAUC.

S2: Multilayer Perceptron implementation details

The implemented Multilayer Perceptron is trained for 25 epochs with a batch size of 1000. The activation function in all but the last layers is the rectified linear unit (ReLU). We use binary crossentropy as the loss function, and assign each observation a weight inversely proportional to its class frequency. We tune the number and size of hidden layers (one to three layers, 100 to 500 neurons per layer, the sum across all layers not exceeding 500), an L1 and L2 penalty (evenly spaced on the log-scale between $10e-5$ and $10e-3$), a dropout fraction (evenly spaced between 0 and 0.3), and the learning rate (evenly spaced on the log-scale between $10e-5$ and $10e-3$).

To model the effect of weight changes on the risk of unplanned HF hospitalisation, the optimal combination of three to 15-day weight differences is chosen via cross-validation during hyperparameter optimization. We add either one or two weight differences between 2 and 15 days. In case of two differences, the smaller is restricted to be less or equal to four, and the two parameters need to be at least 7 apart.

Hyperparameter tuning consists of 50 random draws from the hyperparameter space and 5-fold cross-validation.

Using the optimal choice of hyperparameters, we compute the permutation importance scores per predictor in the same fashion as described above for the Random Forest model (using 5-fold, instead of 10-fold cross-validation). For the final model, we select only predictors with a permutation importance larger than zero, and re-do the hyperparameter tuning with only those.

S3: Merging the baseline risk variable and the model for daily assessment of risk of HF hospitalisation

To make the output of the baseline risk model available for the daily risk calculation without data leakage, we assigned patients from the training set a base risk score using leave-one-out predictions. While the base risk model was trained only on predictors collected during the baseline examination, predictions used in the risk model for unplanned HF hospitalisation were updated based on the latest available information collected during later quarterly examinations performed as part of the trial.