

Supplementary material: OpenBUGS code

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We will describe the OpenBUGS code used to implement the model presented in Figure 8. Code for the models in Figures 2 to 7 can be obtained as simplifications of the code presented here.

Inputs to the OpenBUGS code, such as data and hyperparameters, were prepared using the R software and read into OpenBUGS using the package `R2openBUGS`. In particular, covariate data from both CE-MARC and CECaT were shifted and scaled in R using baseline summaries from the EUROPA study (except for symptomatic angina, which was not an input to either logistic regression). For example, 44% of EUROPA patients were on nitrates at baseline, so the binary covariate for nitrate use was shifted to take values -0.44 (not on nitrates) and 0.56 (on nitrates).

From CE-MARC (Study 2), each available variable was provided to OpenBUGS as a vector of length 272 (one element for each patient), including time to first CV event (`y2`), time of censoring (`c2`), and covariates such as use of nitrates (`nitrates2`) and age over 65 (`age2`). If patient i was censored then we set the i -th event time to be missing (`y2[i] <- NA`). If patient i experienced a CV event then we set the i -th censoring time to be zero, (`c2[i] <- 0`). Similarly, covariate data from CECaT (Study 3) were provided as vectors of length 217 (`nitrates3`, `age3`, etc.).

Hyperparameters describing the prior distribution of the parameter vector β also required attention. The first 17 elements of β were assigned a multivariate normal distribution (with mean and precision as estimated in EUROPA) and we specified an independent normal prior (mean 0 and standard deviation 0.97) for the 18th element, resulting in a multivariate normal prior for β overall (with known mean vector `mu_beta` and precision matrix `Tau_beta`). However,

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it was necessary to reparametrise due to the shifting and scaling of covariates. Again, this was performed in R (requiring only straightforward matrix multiplication), resulting in a multivariate normal distribution with mean vector `mu_beta_reparam` and precision matrix `Tau_beta_reparam` as inputs to OpenBUGS.

The following OpenBUGS code was used to sample (approximately) from the posterior distribution of β . Further manipulations of the OpenBUGS output, such as burn-in, thinning, reparametrisation and diagnostic checks, were performed in R.

```
model{

  # Prior for beta (reparametrised version):

  beta[1:18] ~ dmnorm(mu_beta_reparam[ ], Tau_beta_reparam[ , ])

  # Priors for logistic regression parameters:

  theta[1:11] ~ dmnorm(mu_theta[ ], Tau_theta[ , ])
  gamma[1:12] ~ dmnorm(mu_gamma[ ], Tau_gamma[ , ])

  # Loop over 217 CECaT patients:

  for(j in 1:217){

    # Imputing nitrate use for CECaT patients:
    # s[j] is the probability patient j used nitrates (x* = 0.56),
    # in the logistic regression p(x* | x'', theta).

    logit(s[j]) <- theta[1] +
                   theta[2] * bloodpress3[j] +
                   theta[3] * age3[j] +
                   theta[4] * male3[j] +
                   theta[5] * smoker3[j] +
                   theta[6] * diabetes3[j] +
                   theta[7] * familyhistory3[j] +
                   theta[8] * lipid3[j] +
                   theta[9] * obese3[j] +
                   theta[10] * mi3[j] +
                   theta[11] * stenosis3[j]
```

```
# Use of nitrates is generated from a Bernoulli distribution,
# giving values 0 and 1 which we shift to -0.44 and 0.56:

nitrates3_imp[j] ~ dbern(s[j])
nitrates3[j]     <- nitrates3_imp[j] - 0.44

# Fitting the model for angina:
# q[j] is the probability patient j had angina (x' = 1),
# in the logistic regression p(x' | x'', x*, gamma).

logit(q[j]) <- gamma[1]          +
                gamma[2] * bloodpress3[j] +
                gamma[3] * age3[j]      +
                gamma[4] * male3[j]     +
                gamma[5] * smoker3[j]   +
                gamma[6] * diabetes3[j] +
                gamma[7] * familyhistory3[j] +
                gamma[8] * lipid3[j]    +
                gamma[9] * obese3[j]    +
                gamma[10] * nitrates3[j] +
                gamma[11] * mi3[j]      +
                gamma[12] * stenosis3[j]

angina3[j] ~ dbern(q[j])

} # End of loop over CECaT patients.

# Loop over 272 CE-MARC patients:

for(i in 1:272){

  # Uncertainty in previous MI:
  # Previous acute coronary syndrome (acs2[i] = 0.35) is taken
  # to have probability 44% of being MI (mi2[i] = 0.35). Otherwise
  # (acs2[i] = -0.65) no previous MI was assumed (mi2[i] = -0.65).

  notmi[i] ~ dbern(0.56)
  mi2[i]   <- acs2[i] - notmi[i] * (acs2[i] + 0.65)
```

```
# Fitting the model for nitrate use:
# r[i] is the probability patient i used nitrates (x* = 0.56),
# in the logistic regression p(x* | x'', theta).

logit(r[i]) ~ theta[1] +
              theta[2] * bloodpress2[i] +
              theta[3] * age2[i] +
              theta[4] * male2[i] +
              theta[5] * smoker2[i] +
              theta[6] * diabetes2[i] +
              theta[7] * familyhistory2[i] +
              theta[8] * lipid2[i] +
              theta[9] * obese2[i] +
              theta[10] * mi2[i] +
              theta[11] * stenosis2[i]

# Shifting is required as OpenBUGS requires observations
# from a Bernoulli distribution to be 0 or 1:

nitrates2_shift[i] <- nitrates2[i] + 0.44
nitrates2_shift[i] ~ dbern(r[i])

# Imputing angina for CE-MARC patients:
# p[i] is the probability patient i had angina (x' = 1)
# in the logistic regression p(x' | x'', x*, gamma).

logit(p[i]) <- gamma[1] +
               gamma[2] * bloodpress2[i] +
               gamma[3] * age2[i] +
               gamma[4] * male2[i] +
               gamma[5] * smoker2[i] +
               gamma[6] * diabetes2[i] +
               gamma[7] * familyhistory2[i] +
               gamma[8] * lipid2[i] +
               gamma[9] * obese2[i] +
               gamma[10] * nitrates2[i] +
               gamma[11] * mi2[i] +
               gamma[12] * stenosis2[i]

angina2[i] ~ dbern(p[i])
```

```

# Fitting the model for time to CV event:
# lambda[i] is the CV event rate of patient i in the
# proportional hazards model p(y | x, beta).

log(lambda[i]) <- beta[1] +
  beta[2] * ace2[i] +
  beta[3] * age2[i] +
  beta[4] * male2[i] +
  beta[5] * smoker2[i] +
  beta[6] * tiapvdcva2[i] +
  beta[7] * diabetes2[i] +
  beta[8] * familyhistory2[i] +
  beta[9] * angina2[i] +
  beta[10] * bloodpress2[i] +
  beta[11] * clearance2[i] +
  beta[12] * obese2[i] +
  beta[13] * cholesterol2[i] +
  beta[14] * nitrates2[i] +
  beta[15] * mi2[i] +
  beta[16] * calcium2[i] +
  beta[17] * lipid2[i] +
  beta[18] * stenosis2[i]

# Each event time y2[i] is assumed to come from an
# exponential distribution with rate lambda[i].
# y2[i] was either observed or was censored at c2[i]
# (and therefore sampled assuming y2[i] > c2[i]):

y2[i] ~ dexp(lambda[i])I(c2[i], )

# Posterior predictive checks:
# We predict event times for each patient, using
# the cut function to prevent these from influencing
# the parameter distributions.

lambda_copy[i] <- cut(lambda[i])
y2_pred[i] ~ dexp(lambda_copy[i])

} # End of loop over CE-MARC patients.

} # End of Bayesian model specification.

```