## Predict Prostate Mathematics

## Summary

- Predict takes the form of a competing risk Cox survival model, with fractional polynomial baseline cumulative hazards.
- Approximate intervals for the benefits of treatment can be obtained from the treatment-effects uncertainties.

## The form of the Predict Prostate V1.1 algorithm

The estimated baseline cumulative hazard for prostate cancer mortality  $H_{pc}$  at t days post-diagnosis has the form:

$$H_{\rm pc}(t) = \exp[a_{\rm c}f(t)]$$

where  $a_c$  is a vector of estimated coefficients, and f a (column) vector of fractional polynomial functions of time post-diagnosis.

In Predict Prostate 1.1,

$$H_{pc}(t) = \exp[-16.40532 + 1.653947*(\log(t)) + (1.89x(10^{-12}))*(t^{3})]$$

The estimated survival function for prostate cancer mortality  $S_{pc}$  given prostate cancer risk factors  $\chi_{RP}$  and the i treatment option  $\chi_i$  is given by

$$S_{pc}^{i}(t) = \exp \left[-H_{pc}(t)\right] \exp \left[b' \chi_{RP} + c' \chi_{i}\right]$$

where b and c vectors of estimated coefficients. This is the chance of living beyond *t* days after diagnosis, assuming only prostate cancer mortality.

The estimated baseline cumulative hazard for other-cause mortality  $H_o$  has the form

$$H_o(t) = \exp[a_o'q(t)]$$

where  $a_0$  is a vector of estimated coefficients, and g a vector of fractional polynomial functions of time post diagnosis. This is the chance of living beyond t days after diagnosis, assuming only other-cause mortality.

The estimated survival function for non-prostate cancer (other) mortality  $S_0$ , is given by

$$S_o(t) = \exp \left[-H_o(t) \exp[b_o' \chi_{Ro}]\right]$$

where  $b_o$  is a vector of estimated coefficients, and other cause risk factors is given by  $\chi_{Ro}$ 

In Predict Prostate 1.1, the parameters given by

$$H_o(t) = \exp[-12.4841 + 1.32274*(\log(t)) + 2.90x(10^{-12}))*(t^3)]$$

 $S_o(t)$  is the yellow 'dashed' line in the graphs - the survival of men who are assumed not to die of prostate cancer, essentially who are 'cured'.

The overall estimated survival function S(t), assuming independent competing risks and treatment regime i is given by

$$S^{i}(t) = S_{o}(t)S_{pc}^{i}(t)$$

which is the estimated chance of living beyond time t. This is a competing risk Cox survival model.

Overall survival with conservative management or 'no treatment' (i = 0) is then

$$S^{0}(t) = S_{o}(t)S_{pc}^{0}(t)$$

The overall survival benefit of radical treatment (i = 1) at time t is then given by

$$S^{1}(t) - S^{0}(t) = S_{o}(t)(S_{pc}^{1}(t) - S_{pc}^{0}(t))$$

which is the benefit in prostate cancer specific survival  $S_{\rho c}^{\ 1}(t)$  -  $S_{\rho c}^{\ 0}(t)$ , scaled by the probability of surviving other risks  $S_o(t)$ .