## Nelson-Aalen & Kaplan-Meier

**Basic Relations** 

$$S(t) = \exp(-\int_0^t \alpha(s)ds) \qquad A(t) = \int_0^t \alpha(s)ds \qquad -S'(t) = \alpha(t)S(t) \qquad f(t) = \alpha(t)S(t) \qquad P(T > x|T > y) = \frac{P(T > x)}{P(T > y)} = \frac{S(x)}{S(y)}$$

**Nelson-Aalen** Non-parametric estimator of the cumulative hazard rate.  $A(t) = \int_0^t \alpha(u) du$  where  $\alpha(t)$  is the hazard rate at time t.

Formulas 
$$\hat{A}(t) = \sum_{T_i < t} \Delta \hat{A}(T_j)$$

No ties 
$$\Delta \hat{A}(T_j) = \frac{1}{Y(T_j)}$$
 Rounded Ties  $\Delta \hat{A}(T_j) = \sum_{k=1}^{d_j-1} \frac{1}{Y(T_j)-k}$  True Ties  $\Delta \hat{A}(T_j) = \frac{d_j}{Y(T_j)}$   $\hat{\sigma}_{N-A}^2(t) = \sum_{T_j \leq t} \Delta \hat{\sigma}^2(T_j)$ 

No ties 
$$\Delta \hat{\sigma}^2(T_j) = \frac{1}{Y(T_j)^2}$$
 Rounded Ties  $\Delta \hat{\sigma}^2(T_j) = \sum_{k=1}^{d_j-1} \frac{1}{(Y(T_j)-k)^2}$  True Ties  $\Delta \hat{\sigma}^2(T_j) = \frac{(Y(T_j)-d_j)d_j}{Y(T_j)^3}$ 

**Derivation of the Nelson-Aalen Estimator**. M(t) is a mean-zero m.g.  $H(t) = \frac{J(t)}{Y(t)}$  is predictable where  $J(t) = \mathbf{1}(Y(t) > 0)$ . We then get  $\hat{A}(t) = \int_0^t H(s)dN(s) = \int_0^t \alpha(s)J(s)ds + \int_0^t \frac{J(s)}{Y(s)}dM(s)$ . As M(t) is mean-zero m.g. we then get that the Nelson-Aalen estimator is a unbiased estimator of  $A^*(t) = \int_0^t \alpha(s)J(s)ds$  since  $\mathbb{E}\left[\hat{A}(t) - A^*(t)\right] = 0$ .  $\hat{A}(t)$  is however a biased estimator of A(t) since  $\mathbb{E}[J(s)] = \mathbb{P}(Y(s) > 0)$ . This bias is however very small.

**Derivation of variance of Nelson-Aalen Estimator**. Recall that 
$$\left[\int HdM\right](t) = \int H(s)^2 dN(s)$$
 where we have  $H(s) = \frac{J(s)}{Y(s)} \Longrightarrow \left[\int HdM\right](t) = \int \left(\frac{J(s)}{Y(s)}\right)^2 dN(s) = \left[\hat{A} - A^*\right](t) \Longrightarrow \hat{\sigma}_{N-A}^2 = \sum_{T_j \leq t} \frac{1}{Y(T_j)}$ 

Delta Method CI Nelson-Aalen We have that  $\hat{A}(t) \overset{\text{approx}}{\sim} N(A(t), \hat{\sigma}^2(t))$ 

$$g(\hat{A}(t)) \approx g(A(t)) + g'(A(t)) \underbrace{(\hat{A}(t) - A(t))}_{\mathbb{E}[\dots] = 0} \implies \mathbb{E}[g(\hat{A}(t))] \approx g(A(t)) \text{ and } \mathbb{E}[(g(\hat{A}(t)) - g(A(t))^2] \approx g'(\hat{A}(t))^2 \underbrace{\mathbb{E}[(\hat{A}(t) - A(t))^2]}_{\hat{\sigma}^2}$$

$$\implies g(\hat{A}(t)) \overset{\text{approx}}{\sim} N(g(A(t), |g'(\hat{A}(t)|\hat{\sigma})))$$

Let  $g(x) = \log(x)$  which then gives us  $g^{-1}(x) = e^x$  and  $g'(x) = \frac{1}{x}$ . The interval then becomes as follows.

$$g^{-1}(CI) = \exp\left\{\log(\hat{A}(t)) \pm z_{1-\alpha/2} \frac{\hat{\sigma}}{\hat{A}(t)}\right\} = \hat{A}(t) \exp\left\{\pm z_{1-\alpha/2} \frac{\hat{\sigma}}{\hat{A}(t)}\right\}$$

**Kaplan-Meier** Non-parametric estimator of the survival function.  $S(t) = e^{-A(t)}$  where A(t) is the cumulative hazard rate at time t.

Formulas 
$$\hat{S}(t) = \prod_{T_j \le t} \left( 1 - \frac{1}{Y(T_j)} \right) = \prod_{T_j \le t} (1 - \Delta \hat{A}(T_j))$$
  
 $\hat{\tau}^2(t) = \hat{S}(t)^2 \sum_{T_j \le t} \frac{1}{Y(T_j)^2} = \hat{S}(t)^2 \hat{\sigma}_{N-A}^2 \quad \hat{\tau}^2(t) = \hat{S}(t)^2 \sum_{T_j \le t} \frac{d_j}{Y(T_j)(Y(T_j) - d_j)}$  (Greenwood)

**Derivation of the Kaplan-Meier Estimator** Recall  $\mathbb{P}(T > t) \implies S(t_k|t_{k-1}) = \mathbb{P}(T > t_k|T > t_{k-1}) = \frac{S(t_k)}{S(t_{k-1})}$ . Let  $0 = t_0 < t_1 < \ldots < t_n$  and note that  $\mathbb{P}(T > t_0) = 1$  which gives us  $S(t_n) = \prod_{k=1}^n \frac{S(t_k)}{S(t_{k-1})}$ . We formally we define the survival function as  $S(t) = \prod_{u \leq t} (1 - dA(u))$  since  $\frac{S(t_k)}{S(t_{k-1})} = dA(t_k)$  when  $t_k - t_{k-1} < 1$ . This gives us the estimator  $\hat{S}(t) = \prod_{T_j \leq t} (1 - \Delta \hat{A}(T_j))$  as  $\Delta \hat{A}(t)$  serves as an estimator for dA(t).

**Kaplan Meier CI**  $\hat{S}(t) \pm z_{1-\alpha/2}\hat{\tau}(t)$ . Log-transforms (using same method as for Nelson-Aalen above) etc.

**Derivation of variance of Kaplan-Meier Estimator** Let  $S^*(t) = \iint_{u \le t} (1 - dA^*(t))$  where  $A^*(t) = \int_0^t J(u) dA(u)$ . If  $\mathbb{P}(J(s) = 0) << 1$  then  $S^*$  and S are close. We measure this closeness by  $\frac{\hat{S}(t)}{S^*(t)} - 1 = -\int_0^t \frac{\hat{S}(u-)}{S^*(u)} d(\hat{A} - A^*)(u)$ . We then have that  $\mathbb{E}\left[\frac{\hat{S}(t)}{S^*(t)}\right] = 1$ . We can

then repat the arguments as we do for the variance of the Nelson-Aalen estimator above. 
$$\left[\frac{\hat{S}}{S^*} - 1\right] = \left[\int \frac{-\hat{S}}{S^*} \underbrace{d(\hat{A} - A^*)}_{dM}\right] = \{\text{Theorem}\} = \{0\}$$

 $\int \left(\frac{\hat{S}}{S^*}\right)^2 d[M]. \text{ Note that } M \text{ is the same mean-zero m.g. as in the Nelson-Aalen case which gives us } d[M](t) = \frac{J(t)}{Y(t)} dN(t). \text{ This does in turn give us that } \left[\frac{\hat{S}}{S^*} - 1\right] = \int \left(\frac{\hat{S}}{S^*}\right)^2 \frac{J}{Y^2} dN. \text{ Now by assuming } S^* = S \text{ and } \hat{S}(u) \approx \hat{S}(u-) \text{ we get that } \text{Var}\left(\frac{\hat{S}(t)}{S(t)} - 1\right) = \hat{\sigma}_{\hat{S}/S-1}^2(t) = \int_0^t \frac{J}{Y^2} dN = \hat{\sigma}_{\text{N-A}}^2(t) \implies \hat{\sigma}_{\hat{S}}^2(t) \approx \hat{S}^2(t) \int_0^t \frac{J}{Y^2} dN = \hat{S}^2(t) \hat{\sigma}_{\text{N-A}}^2(t)$ 

## Martingales

**Definition of Martingales** M is a Martingale if  $E[M_t|\mathcal{F}_s] = M_s$ ,  $t \geq s$  and  $E[|M_t|] < \infty$ .

**Formulas** 

$$(H \bullet M)_n = H_0 M_0 + H_1 (M_1 - M_0) + \dots + H_n (M_n - M_{n-1})$$

$$\langle H \bullet M \rangle_n = \sum_{i=1}^n H_i^2 \Delta \langle M \rangle_i \text{ where } \Delta \langle M \rangle_i = [(M_i - M_{i-1})^2 | \mathcal{F}_{i-1}]$$

$$\langle H \bullet M \rangle = H^2 \bullet \langle M \rangle$$

$$\langle M \rangle_n = \sum_{i=1}^n \text{Var}(\Delta M_i | \mathcal{F}_{i-1}) = \sum_{i=1}^n \mathbb{E}[(M_i - M_{i-1})^2 | \mathcal{F}_{i-1}]$$

$$\langle \int H dM \rangle = \int H^2(s) d\langle M \rangle(s) = \int H^2(s) \lambda(s) ds$$

$$Cov(M_s, M_t - M_s) = 0$$

$$M^2 - \langle M \rangle \text{ and } M^2 - [M] \text{ are zero mean m.g.s.}$$

$$Var(M(t)) = E(M(t))^2 = E\langle M \rangle(t) = E[M](t)$$

$$\Delta M_t = M_t - M_{t-1} \text{ called m.g. difference}$$

$$[H \bullet M]_n = \sum_{i=1}^n H_i^2 \Delta[M]_i \text{ where } \Delta[M]_i = (M_i - M_{i-1})^2$$

$$[H \bullet M] = H^2 \bullet [M]$$

$$[M]_n = \sum_{i=1}^n (\Delta M_i)^2 = \sum_{i=1}^n (M_i - M_{i-1})^2$$

$$\left[\int H dM\right] = \int H^2(s) d[M]s) = \int H^2(s) dN(s)$$

$$E[M_t - M_s | \mathcal{F}_s] = E[M_t | \mathcal{F}_s] - M_s = 0$$

$$E[\Delta M_t | \mathcal{F}_{t-1}] = 0$$

**Doob decomposition** Let X with  $X_0$  be a general discrete time proc. and let M be def. by  $M_0 = X_0 = 0$  and  $M_n - M_{n-1} = X_n - E[X_n|\mathcal{F}_{n-1}] \implies E[\Delta M_n|\mathcal{F}_{n-1}] = 0 \implies X_n = E[X_n|\mathcal{F}_{n-1}] + \Delta M_n = \underbrace{E[X_n|\mathcal{F}_{n-1}]}_{pred} + \underbrace{(X_n - E[X_n|\mathcal{F}_{n-1}])}_{noise}$ 

## Frailty

Basic Relations  $A(t|Z) = \int_0^t \alpha(s|Z)ds$  if proportional frailty (i.e.  $\alpha(t|Z) = \alpha(t)Z$  we have  $A(t|Z) = ZA(t) \implies S(t) = \exp\{-ZA(t)\}$ . Consequently the piopulation survival is given by  $S(t) = \mathbb{E}[S(t|Z)] = \mathbb{E}[\exp\{-A(t|Z)\}] = \mathcal{L}_Z(A(t))$  where  $\mathcal{L}_Z(c)$  is the Laplace transform (i.e.  $\Psi_Z(-c)$ ).

**Population Hazard Rate**  $\mu(t) = \frac{-S'(t)}{S(t)} = \alpha(t) \frac{\mathcal{L}'(A(t))}{\mathcal{L}(A(t))}$  assuming proportional frailty  $(\alpha(t|Z) = \alpha(t)Z)$ 

Modelling Frailty as Power Variance Function We let  $Z \sim \text{PVF}(\varphi, \nu, m)$  for  $\nu, m+1, m\varphi > 0$ . This gives us  $\mathbb{E}[Z] = \frac{\varphi m}{\nu}$  and  $\text{Var}(Z) = \frac{\varphi m}{\nu} \frac{m+1}{\nu}$ . For a PVF it holds that  $S(t) = \exp\left\{-\varphi\left(1 - \left(\frac{1}{1 + \frac{A(t)}{\nu}}\right)^2\right)\right\}$  and  $\mu(t) = \frac{\varphi m}{\nu} \frac{\alpha(t)}{(1 + \frac{A(t)}{\nu})^{m+1}}$ .

Modelling Frailty as a Gamma distribution We have that  $Z \sim \Gamma(\nu, \eta), \nu, \eta > 0$ . We then have  $\mathbb{E}[Z]$  and  $\Phi_Z(c) = \left(\frac{\nu}{\nu-c}\right)^{\eta}$  (see Distributions on page 6). This gives us that  $\mathcal{L}_Z(c) = \frac{1}{\left(1+\frac{c}{\nu}\right)^{\eta}}$ . Common to use  $\mathbb{E}[Z] = 1 \implies \nu = \eta$ . This then gives us  $\mathcal{L}_Z(c) = \frac{1}{\left(1+\frac{c}{\nu}\right)}$ . If we let  $\delta = \frac{1}{\nu} = \text{Var}(Z)$  which implies  $\mathcal{L}_Z(c) = \frac{1}{(1+\delta c)^{1/\delta}}$ . Finally (if we assume a proportional frailty model) we get that the population survival  $S(t) = \mathcal{L}_Z(A(t)) = \mathcal{L}_Z(c) = \frac{1}{(1+\delta A(t))^{1/\delta}}$  and the population hazard  $\mu(t) = \frac{-S'(t)}{S(t)} = \frac{\alpha(t)}{1+\delta A(t)}$ 

# **Testing**

**Breslow-Estimator**  $\hat{A}_0(t) = \hat{A}_0(t; \hat{\beta})$  where  $\hat{A}_0(t; \hat{\beta}) = \int_0^t \frac{dN_{\bullet}(u)}{\sum_{l=1}^n Y_l(u)r(\beta, \mathbf{x}_l)}$ , Cox prop. haz. model  $\implies \alpha(t|\mathbf{x}) = \alpha_0(t)r(\beta\mathbf{x}) \implies A(t|\mathbf{x}) = A_0(t)r(\beta\mathbf{x})$ 

**Gehan-Breslow Test**  $U(t_0) = \frac{Z_1(t_0)}{\sqrt{V_{11}}} \sim N(0,1)$  where  $Z_1(t_0) = \int_0^{t_0} Y_2(t) dN_1(t) - \int_0^{t_0} Y_1(t) dN_2(t)$  and  $V_{11}(t_0) = \int_0^{t_0} Y_1(t) Y_2(t) dN_{\bullet}(t)$ 

Cox regression Note that  $r(\boldsymbol{\beta}, \mathbf{x}) = \boldsymbol{\beta}^T \mathbf{x}$ .

Multiplicative model:  $\alpha(t|x_1,...,x_p) = \alpha_0(t) \exp(\beta_1 x_1 + ... + \beta_p x_p)$ . Additive model:  $\alpha(t|x_1,...,x_p) = \underbrace{\beta_0(t)}_{baselinehaz} + \beta_1 x_1 + .... + \beta_p x_p$ . Cox partial likelihood:  $L(\beta) = \prod_{T_j} \frac{r(\beta,x_j)}{\sum_{i \in \mathcal{R}_j} r(\beta,x_j)}$  where  $\mathcal{R}_j$  is the risk set just before event j.

LR test:  $\chi_{LR}^2 = 2(\log(L(\hat{\beta}) - \log(\beta_0)) \sim \chi^2(1)$ 

 $\textbf{Hazard Ratio} \ \frac{\alpha(t|\mathbf{x}_1)}{\alpha(t|\mathbf{x}_2)} \ \text{e.g.} \ \frac{\alpha(t|\mathbf{x}=(1,0))}{\alpha(t|\mathbf{x}=(0,1))} = \frac{r(\beta,(1,0))}{r(\beta,(0,1))}$ 

#### Misc

Accelerated failure time models:  $\log U_i = \beta^T \mathbf{x}_i + \varepsilon_i$  where  $E[\varepsilon_i] = 0$  iid.  $\Longrightarrow S_{U_i}(u) = P(U_i > u) = P(e^{\beta^T \mathbf{x}_i + \varepsilon_i} > u) = P(\underbrace{\varepsilon_i} > ue^{-\beta^T \mathbf{x}_i} = S_{w_i}(ue^{-\beta^T \mathbf{x}_i}), \text{ (change of time)}, \implies S'_{U_i}(u) = S'_{w_i}(ue^{-\beta^T \mathbf{x}_i})e^{-\beta^T \mathbf{x}_i} \implies \alpha_{U_i}(u) = \frac{-S'_{U_i}(u)}{S_{U_i}(u)} = \alpha_{w_i}(ue^{-\beta^T \mathbf{x}_i})ue^{-\beta^T \mathbf{x}_i}$ 

**Likelihood** With censored observations we can express the likelihood as follows.  $L(\theta; t_1, ..., t_n) \prod_{i=1}^n \mathbb{P}(T = t_i)_i^{\delta} \mathbb{P}(T \ge t_i)^{1-\delta_i}$  where  $\delta_i$  is the indicator of  $t_i$  being censored. See example below under **Examples** 

Likelihood in terms of hazard rate  $L(\theta) = \prod_{i \in \mathcal{D}_i} \mathbb{P}(T \in [t_i, t_i + dt; \theta)) \prod_{i \notin \mathcal{D}_i} \mathbb{P}(T \geq t_i; \theta) \approx \prod_{i \in \mathcal{D}_i} f(t_i; \theta) dt \prod_{i \notin \mathcal{D}} S(t_i; \theta) \propto \prod_{i \in \mathcal{D}_i} \alpha(t_i; \theta) S(t_i; \theta) \prod_{i \notin \mathcal{D}_i} S(t_i; \theta) = \prod_{i=1}^n \alpha(t_i, \theta)^{\delta_i} S(t_i; \theta)$  where individuals with  $\delta_i = 1$  belong to  $\mathcal{D}_i$ .

### Problems, solutions and examples

 $\begin{aligned} \mathbf{Problem \ 1a \ Show} \ \hat{A}(t) &= \int_0^t \frac{I(Y(s)>0)}{Y(s)} dN(s) \text{ is an unbiased estimator of } A^*(t) = \int_0^t I(Y(s)>0) \alpha(s) ds. \\ \mathbf{Solution} \ \hat{A}(t) &= A^*(t) = \int_0^t \frac{I(Y(s)>0)}{Y(s)} dN(s) - \int_0^t I(Y(s)>0) \alpha(s) ds = \int_0^t \frac{I(Y(s)>0)}{Y(s)} dN(s) - \int_0^t \frac{I(Y(s)>0)}{Y(s)} Y(s) \alpha(s) ds = \int_0^t \frac{I(Y(s)>0)}{Y(s)} dN(s) - \int_0^t \frac{I(Y(s)>0)}{Y(s)} Y(s) \alpha(s) ds = \int_0^t \frac{I(Y(s)>0)}{Y(s)} dN(s) - Y(s) \alpha(s) ds = \int_0^t \frac{I(Y(s)>0)}{Y(s)} \frac{dM(s)}{Y(s)} dN(s) - I(s) \alpha(s) ds = \int_0^t \frac{I(Y(s)>0)}{Y(s)} dN(s) ds = \int_0^t \frac{I(Y(s)>0)$ 

**Problem 1b** Show  $A^*(t)$  is a biased estimator of  $A(t) = \int_0^t \alpha(s) ds$ .

**Solution**  $E[A^*(t)] = E[\int_0^t I(Y(s) > 0)\alpha(s)ds] \le E[\int_0^t 1 \cdot \alpha(s)ds] = A(t).$ 

**Problem 1c** Calculate the optional variation of  $\hat{A}(t)A^*(t)$ , i.e.  $[\hat{A}(t)A^*(t)]$  and write this expression as a sum.

Solution  $[\hat{A}(t)A^*(t)] = [\int \frac{1}{Y} dM](t) = \int_0^t \left(\frac{I(Y(s)>0)}{Y(s)}\right)^2 dN(s) = \int_0^t \frac{I(Y(s)>0)}{Y(s)^2} dN(s).$ 

We also have that  $Var(\hat{A}(t) - A^*(t)) = E([\hat{A}(t) - A^*(t)](t))$  and thus  $\hat{A}(t) - A^*(t)](t)$  is an unbiased est. of the variance.

Morover, assuming no ties  $[\hat{A}(t) - A^*(t)](t) = \int_0^t \frac{I(Y(s)>0)}{Y(s)^2} dN(s) = \sum_{T_j \le t} \frac{1}{Y(T_j)}$ , which is the Nelson-Aalen estimator.

**Problem 2** Let  $X_n$  be discrete time m.g. Show that  $E[X_n^2]$  is non-decreasing in n.

Solution First show  $M_{n+1} = X_n(X_{n+1} - X_n)$  has zero mean.  $E[X_n(X_{n+1} - X_n)|\mathcal{F}_n] = X_nE[X_{n+1} - X_n|\mathcal{F}_n] = 0$ . Now note that  $(X_{n+1} - X_n)^2 = (X_{n+1} - X_n)(X_{n+1} - X_n) = X_{n+1}(X_n + 1 - X_n) - M_n$ . We get that  $E[(X_{n+1} - X_n)^2] = E[X_{n+1}(X_n + 1 - X_n) - M_n] = E[X_{n+1}(X_n + 1 - X_n)] = E[X_{n+1}^2] - E[X_{n+1}(X_n + 1 - X_n)] = E[X_n(X_n + 1 - X_n)]$ 

**Problem 3** Let  $T_1, ..., T_n$  i.i.d.  $\operatorname{Exp}(\nu)$ . Let  $c_1, ..., c_n$  be non-random censoring times and  $\tilde{T}_i = \min(T_i, c_i)$ . Let  $D_i = 1$  if  $\tilde{T}_i = T_i$ . Construct the likelihood for this situation.

**Solution** Contribution for an observed event is  $\alpha(\tilde{t}_i;\nu) \exp\{\int_0^{\tilde{t}_i} \alpha(s;\nu) ds\} = \nu \exp\{-\tilde{t}_i\nu\}$ , since  $\alpha(t;\nu) = \nu$ . A censored event contributes to the likelihood with  $S(\tilde{t}_i;\nu) = \exp\{\int_0^{\tilde{t}_i} \alpha(s;\nu) ds\} = \exp\{-\tilde{t}_i\nu\}$ .

By combining these we get  $L(\nu) = \prod_{i=1}^{n} (\alpha(\tilde{t}_i; \nu) \exp\{\int_0^{\tilde{t}_i} \alpha(s; \nu) ds\})^{D_i} (\exp\{\int_0^{\tilde{t}_i} \alpha(s; \nu) ds\})^{1-D_i} = \nu^d \exp\{-\nu r\}$ , where  $d = \sum_i D_i$  and  $r = \sum_i \tilde{t}_i$ .

**Problem 4a** Let N(t) be a Poisson process with intensity function  $\lambda(t)$ . Show that  $M(t) = N(t) - \int_0^t \lambda(s) ds$  is a mean zero m.g.

**Solution** From def. of the Po-process we know  $E[N(t) - N(s)|\mathcal{F}_s] = \int_s^t \lambda(s)ds$  which follows from indep. increments property. Thus,  $E[N(t) - \int_0^t \lambda(s)ds|\mathcal{F}_s] = E[N(t) - \int_0^t \lambda(s)ds - N(s) + N(s)|\mathcal{F}_s] = E[N(t) - N(s)] - \int_0^t \lambda(s)ds + N(s) = \int_s^t \lambda(s)ds - \int_0^t \lambda(s)ds + N(s) = N(s) - \int_0^s \lambda(s)ds$ .

**Problem 4b** For M(t) above, it holds that  $M(t)^2 - \int_0^t \lambda(s) ds$  is a mean-zero m.g. Use this with a) to show  $\lim_{h\to 0^+} \frac{1}{h} E[(M(t+h) - M(t))^2 | \mathcal{F}_t] = \lambda(t)$ . (i.e.  $d\langle M \rangle(t) = \lambda(t)$ .)

Solution  $E[(M(t+h)-M(t))^2|\mathcal{F}_t] = E[(M(t+h)^2|\mathcal{F}_t] - 2E[M(t+h)|\mathcal{F}_t]M(t) + M(t)^2 = E[(M(t+h)^2|\mathcal{F}_t] - M(t).$  Now using that  $E[(M(t+h)^2-\int_0^{t+h}\lambda(u)du|\mathcal{F}_t] = M(t)^2-\int_0^t\lambda(u)du$  it follows that  $E[(M(t+h)-M(t))^2|\mathcal{F}_t] = M(t)^2-\int_0^t\lambda(u)du + \int_0^{t+h}\lambda(u)du - M(t)^2 = \int_t^{t+h}\lambda(u)du$ . Desired result follows from standard calculus.

## Examples

Example of Nelso-Aalen Calculations (different types of ties,  $A_1$  uses true ties whilst  $A_2$  uses rounded ties)

| t   | Y(t) | d(t) | $\Delta \hat{A}_1(t)$ | $\Delta \hat{\sigma}_1^2(t)$ | $\Delta \hat{A}_2(t)$                        | $\Delta \hat{\sigma}_2^2(t)$                       |
|-----|------|------|-----------------------|------------------------------|--|--|
| 0.2 | 16   | 1    | $\frac{1}{16}$        | $\frac{(16-1)\cdot 1}{16^3}$ | $\frac{1}{16}$                               | $\frac{1}{16^2}$                                   |
| 0.5 | 15   | 3    | $\frac{3}{15}$        | $\frac{(15-3)\cdot 3}{15^3}$ | $\frac{1}{15} + \frac{1}{14} + \frac{1}{13}$ | $\frac{1}{15^2} + \frac{1}{14^2} + \frac{1}{13^2}$ |
| 0.7 | 12   | 1    | $\frac{1}{12}$        | $\frac{(12-1)\cdot 1}{12^3}$ | $\frac{1}{12}$                               | $\frac{1}{12^2}$                                   |
| 1.1 | 11   | 1    | $\frac{1}{11}$        | $\frac{(11-1)\cdot 1}{11^3}$ | $\frac{1}{11}$                               | $\frac{1}{11^2}$                                   |

#### **Example Population Hazard Rate**

Let  $\alpha(t|Z) = \alpha(t)Z$  where  $\mathcal{L}_Z(s) = \mathbb{E}[\exp\{-sZ\}] = (1+\delta s)^{-1/\delta}$ . We get that  $\mathcal{L}'(s)\big|_{s=0} = -\mathbb{E}[Z] = -\frac{\mathcal{L}_Z(s)}{1+\delta}\big|_{s=0} = -1 \implies \mathbb{E}[Z] = 1$  by derivating both sides with regards to s. We can obtain the population hazard rate  $\mu(t) = \alpha(t)\frac{-\mathcal{L}_Z'(A(t))}{\mathcal{L}_Z(A(t))} = \frac{\alpha(t)}{1+\delta A(t)}$ . By assuming some form of  $\alpha(t)$  (or A(t)) we can then examine the population hazard rate as  $t \to \infty$ .

**Example of Likelihood Derivations** We have  $(i, t_i, \delta_i) = \{(1, 1.23, 1), (2, 1.97, 1), (3, 1.17, 0)\}$ . If we assume that the times are  $\text{Exp}(\nu)$  then we get  $L(\nu; t_1, t_2, t_3) = \nu^2 e^{-\nu(t_1 + t_2)} e^{-t_3}$ .

Example of LR-test using Cox partial likelihood We have  $\alpha(t;x) = \alpha_0(t)e^{\beta x}$  and  $\{(T_i, \delta_i, x_i)\}_{i=1...5} = \{(1,1,1), (3,1,0), (4,0,1), (7,0,0)\}$  then  $L(\beta) = \frac{e^{\beta}}{e^{\beta}+1+e^{\beta}+1+e^{\beta}} \cdot \frac{1}{1+e^{\beta}+1+e^{\beta}} \cdot \frac{e^{\beta}}{e^{\beta}}$ . We then perform the likelihood ratio test as described under **Testing** using  $\hat{\beta}$  (which we obtain using regular likelihood theory) and  $\beta_0$ .