# ON TESTS FOR SOME STATISTICAL HYPOTHESES FOR DEPENDENT COMPETING RISKS

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#### Introduction

Competing risks data: T = failure time  $\delta =$  an indicator of the failure type,  $\delta \in \{1, 2, ..., k\}$ 

Planning studies: T could be the time to first employment and  $\delta$  indicating the employment obtained.

Acturial / Medical / engineering studies

Sub-distribution function :

$$F(i,t) = P[T \le t, \delta = i],$$

Sub-survival function :

 $S(i,t) = P[T > t, \delta = i],$ 

Sub-density function f(i, t), i = 1, 2, ..., k.

The distribution function of T:  $H(t) = \sum_{i=1}^{k} F(i, t)$ ,

the survivor function of T:  $S(t) = \sum_{i=1}^{k} S(i, t)$ ,

the density function of T is  $h(t) = \sum_{i=1}^{k} f(i, t)$ ,

the probability of failure due to the ith risk  $p_i=F(i,\infty),$ 

cause-specific hazard rate is  $\lambda_i(t) = \frac{f(i,t)}{S(t)}$ .

The latent failure time model:

 $X_1, X_2, \ldots, X_k$  are latent failure times of any unit exposed to k risks.

Observe  $T = \min(X_1, X_2, \ldots, X_k)$ ,

 $\delta = j$  if  $X_j = \min X_i, 1 \le i \le j$ .

Under independence the marginal and hence the joint distribution is identifiable from the probability distribution of the observable random variables  $(T, \delta)$ .

The independence or otherwise of the latent lifetimes  $(X_1, X_2, \ldots, X_k)$  cannot be statistically tested from any data collected on  $(T, \delta)$  and has to be assumed on the basis of a priori information, if any.

The marginal distribution functions  $F_i(x)$  of  $X_i$ may not represent the probability distribution of lifetimes in any practical situation.

Elimination of *jth* risk may change the environment in such a way that  $F_i(x)$  does not represent the lifetime of  $X_i$  in the changed scenario.

Locally Most Powerful Rank Tests

Suppose k = 2, that is , a unit is exposed to two risks of failure denoted by 1 and 0.

*n* units are put to trial , the data consists of  $(T_i, \delta^*_i), i = 1, ..., n$  where  $\delta^* = 2 - \delta$ .

Test the hypothesis  $H_0$ : F(1,t) = F(2,t), for all t.

Under  $H_0$  the two risks are equally effective. However under the alternative hypothesis one would expect that the two risks are not equally effective atleast at some ages.

The likelihood function for n units is given by (see , Aras and Deshpande (1992))

 $L(\underline{T}, \underline{\delta^*}) = \prod_{i=1}^n [f(1, t_i)]^{\delta^*_i} [f(2, t_i)]^{1 - \delta^*_i}$ 

If F(i,t) depends upon the parameter  $\theta$ , then inference about it can be based on the above likelihood function. When T and  $\delta^*$  are independent then Deshpande (1990) proposed the model

 $F(1,t) = \theta H(t), F(2,t) = (1-\theta)H(t).$ 

 $\theta = P[\delta^* = 1]$ , and the likelihood reduces to

$$L(\theta, H) = \theta^{\sum_{i=1}^{n} \delta^{*_{i}}} (1-\theta)^{\sum_{i=1}^{n} (1-\delta^{*_{i}})} \prod_{i=1}^{n} h(t_{i}).$$

The hypothesis F(1,t) = F(2,t) reduces to testing that  $\theta = 1/2$ .

Then the obvious statistic is the sign statistic  $U_1 = \frac{1}{n} \sum_{i=1}^n \delta^*_i$ .

 $nU_1$  has  $B(n,\theta)$  distribution and one can have optimal estimation and testing procedures based on it.

However, if F(1,t) and F(2,t) depend on a parameter  $\theta$  in a more complicated manner then one needs to look at locally most powerful rank tests.

Let  $f(1,t) = f(t,\theta), f(2,t) = h(t) - f(t,\theta)$ 

h(t) and  $f(t,\theta)$  are known density functions and incidence density such that  $f(t,\theta_0) = \frac{1}{2}h(t)$ .

Let  $T_{(1)} \leq T_{(2)} \leq \ldots \leq T_{(n)}$  denote the ordered failure times.

 $W_i = \begin{cases} 1 & \text{if } T_{(i)} \text{ corresponds to first risk} \\ 0 & \text{otherwise.} \end{cases}$ 

Let  $R_j$  be the rank of  $T_j$  among  $T_1, \ldots, T_n$  .

The likelihood of ranks is given by

$$\int_{0 < t_1 < ... < t_n < \infty} \prod_{i=1}^n [f(t_i, \theta)]^{w_i} [h(t_i) - f(t_i, \theta)]^{1 - w_i} dt_i$$

**Theorem :** If  $f'(t, \theta)$  is the derivative of  $f(t, \theta)$ with respect to  $\theta$ , then the locally most powerful rank test for  $H_0: \theta = \theta_0$  against  $H_1: \theta > \theta_0$ is given by: reject  $H_0$  for large values of

$$L_c = \sum_{i=1}^n w_i a_i, \text{ where}$$
$$a_i = \int \dots \int_{0 < t_1 < \dots < t_n < \infty} \frac{f'(t_i, \theta_0)}{f(t_i, \theta_0)} \prod_{i=1}^n [f(t_i, \theta_0) dt_i].$$

Special cases

(i) If the independence model holds with  $\theta_0 = 1/2$ , then sign test is the LMPR test.

(ii) If  $f(1,t) = \frac{1}{2}g(t,0)$  and  $f(2,t) = \frac{1}{2}g(t,\theta), \theta > 0$  and  $g(t,\theta)$  is the logistic density function  $g(t,\theta) = \frac{e^{(x-\theta)}}{[1+e^{(x-\theta)}]^2}$ , then the LMPR test is based on the statistic

$$W^+ = \sum_{i=1}^n W_i R_i.$$

(iii) In case of Lehmann type alternative defined by  $F(1,t) = \left[\frac{H(t)}{2}\right]^{\theta}$ ,  $F(2,t) = H(t) - \left[\frac{H(t)}{2}\right]^{\theta}$ , LMPR test is based on scores  $a_i = E(E_{(j)})$  where  $E_{(j)}$  is the *jth* order statistic from a random sample of size *n* from standard exponential distribution.

For more complicated families of distributions, e.g., Gumbel (1960), the scores are complicated and need to be solved using numerical integration (see Aras and Deshpande (1992)). Tests for bivariate symmetry

Suppose the latent failure times X and Y are dependent.

Their joint distribution is given by F(x, y).

On the basis of independent pairs  $(T_i, \delta^*_i)$  we want to test whether the forces of two risks are equivalent against the alternative that the force of one risk is greater than that of the other.

$$H_0: F(x,y) = F(y,x)$$
 for every  $(x,y)$ 

Theorem : Under the null hypothesis of bivariate symmetry we have

(i)F(1,t) = F(2,t) for all t,

(ii)S(1,t) = S(2,t) for all t,

(iii)  $\lambda(1,t) = \lambda(2,t)$  for all t,

(iv)  $P[\delta^* = 1] = P[\delta^* = 0]$ 

(v) T and  $\delta^*$  are independent.

 $H_1:\lambda(1,t)<\lambda(2,t)$ 

 $H_2$ : F(1,t) < F(2,t)

 $H_3: S(1,t) > S(2,t).$ 

All these alternatives say that risk II is more potent than risk I at all ages t in some stochastic sense.

Sen (1979) considered fixed sample and sequential tests for the null hypothesis of bivariate symmetry of the joint distribution of (X,Y). The alternatives are expressed in terms of  $\pi_1(t) = Pr[\tilde{\delta} = 1 | T = t]$ ,

the conditional probability that the failure is due to first risk , given that failure occurs at time t.

He derived optimal score statistics for such parametric situations. But the statistics cannot be used without the knowledge of the joint distribution F(x, y).

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For testing  $H_0$  against  $H_1$  consider

$$\psi(t) = F(1,t) - F(2,t)$$

$$\int_0^t S(t) [\lambda(1, u) - \lambda(2, u)] du.$$

 $H_1$  holds iff the above function is non-increasing on t.

Consider the following measure of deviation between  $H_0$  and  $H_1$ ,

$$\Delta = \int_{0 < x < y < \infty} [\psi(x) - \psi(y)] \, dF(x) \, dF(y)$$

Its empirical estimator is  $\Delta_n$ 

$$\Delta_n = \int_{0 < x < y < \infty} [\psi_n(x) - \psi_n(y)] \, dF_n(x) \, dF_n(y)$$

where

$$F_{1n}(t) = \frac{1}{n} \sum_{j=1}^{n} I\{\delta_j = 1, T_j \le t\},\$$

$$F_n(t) = \frac{1}{n} \sum_{j=1}^n I\{T_j \le t\},$$

$$\psi_n(t) = 2F_{1n}(t) - F_n(t),$$

are the empirical estimators of  $F_1, F$  and  $\psi$ , respectively.

Then we can write

$$\Delta_n = \frac{1}{n^3} \left[ \frac{n(n^2 - 1)}{6} - 2\sum_{i=1}^n (i - 1)(n - i + 1)W_i \right].$$

Under  $H_0$ ,  $\Delta = 0$ , under  $H_1$ ,  $\Delta > 0$ .

Large values of the statistic are significant. Rejecting  $H_0$  for large values of  $\Delta_n$  is equivalent to rejecting it for small values of the statistic

$$U_2 = \sum_{i=1}^{n-1} i(n-i)W_{i+1}.$$

T and  $\delta$  are independent under  $H_0$ .

Therefore  $W_1, W_2, \ldots, W_n$  are i.i.d. Bernoulli random variables with

$$P\{W_i = 1\} = P\{W_i = 0\} = \frac{1}{2}, i = 1, 2, \dots, n.$$

The moment generating function of the null distribution of  $U_2$  is given by

$$M(t) = 2^{-n+1} \prod_{i=1}^{n-1} (1 + e^{a_i t}),$$

where  $a_i = i(n - i), i = 1, 2, \dots, n - 1$ .

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For n = 5(1)20, the 5% and 1% critical values of  $U_2$  are given below.

Critical points of  $U_2$  and exact significance levels.

n	lphapprox 0.01	lphapprox 0.05
5	-	0, 0.0625
6	-	0, 0.03125
7	0, 0.015625	6, 0.046875
8	7, 0.015625	14, 0.046875
9	8, 0.011719	22, 0.054688
10	18, 0.011719	34, 0.054688
11	28, 0.013672	50, 0.050781
12	38, 0.010254	67, 0.051758
13	54, 0.010498	90, 0.051514
14	74, 0.01001	118, 0.052734
15	98, 0.010132	148, 0.050293
16	126, 0.01001	186, 0.0513
17	158, 0.01001	228, 0.050308
18	198, 0.010269	278, 0.050613
19	242, 0.010311	332, 0.049911
20	290, 0.01005	396, 0.050467

$$E[U_2] = \frac{n(n^2 - 1)}{12}, \quad Var[U_2] = \frac{n(n^4 - 1)}{120}$$

By the Central Limit Theorem, it can be shown that, under  $H_0$ ,

$$n^{\frac{1}{2}}\left\{\frac{U_2}{n^3} - \frac{1}{12}\right\} \xrightarrow{\mathcal{L}} N(0, \frac{1}{120}).$$

To give an idea about the accuracy of the normal approximation of  $U_2$ , we give the exact (respectively, approximate) significance levels for n = 10 and n = 20.

For n = 10,  $P[U_2 \le 18] = 0.011719$  (resp. 0.0118) and  $P[U_2 \le 34] = 0.054688$  (resp. 0.0594).

For n = 20,  $P[U_2 \le 290] = 0.01005$  (resp. 0.0111) and  $P[U_2 \le 396] = 0.050467$  (resp. 0.049).

Deshpande (1990) proposed two tests for testing  $H_0$  versus  $H_2$ : F(1,t) < F(2,t) on heuristic grounds.

The first test is the Wilcoxon signed rank type statistic

 $W^+ = \sum_{i=1}^n (1 - \delta^*_i) R_i.$ 

It was felt that  $W^+$  will be large when the alternative  $H_2$  is true, there being a greater incidence of the second risk upto any fixed time t.

Another test is based on the U-statistic

$$U_{3} = \frac{1}{\binom{n}{2}} \sum_{1 \le i < j \le n} \phi_{3}(T_{i}, \delta^{*}_{i}, T_{j}, \delta^{*}_{j})$$

$$\phi_{3}(T_{i}, \delta^{*}_{i}, T_{j}, \delta^{*}_{j}) = \begin{cases} 1 & \text{if } \delta^{*}_{j} = 0, T_{i} > T_{j} \\ \text{or } \delta^{*}_{i} = 0, T_{i} < T_{j} \\ 0 & \text{otherwise} \end{cases}$$

The kernel  $\phi_3$  takes value 1 if , and only if, a Y observation is the smallest among  $(X_i, X_j, Y_i, Y_j)$ .

$$\binom{n}{2}U_3 = \sum_{i=1}^n (n - R_i + 1)\delta_i^*.$$

 $E(U_3) = 1/2$  under  $H_0$  and strictly larger than 1/2 under  $H_2$ .

 $U_3$  is same as the statistic proposed earlier to test for  $H_0$  against the alternative  $H_{A1}$  of stochastic dominance of distribution functions of independent latent failure times (see, Deshpande and Dewan (2003)) is also consistent for testing bivariate symmetry against dominance of incidence functions. For testing  $H_0$  against  $H_2$ , one can consider the measure of deviation F(2,t)-F(1,t), which is non-negative under  $H_2$ .

 $\int_0^\infty [F(2,t) - F(1,t)] dH(t)$  $= P[\delta^*_1 = 0, T_1 \le T_2] - \frac{1}{2}.$ 

A U-statistic estimator of this parameter is the statistic  $U_3$  discussed above.

Similarly for testing  $H_0$  against  $H_3$  consider the measure of deviation S(1,t) - S(2,t), which is non-negative under  $H_3$ . Then

 $\int_0^\infty [S(1,t) - S(2,t)] dH(t)$  $= P[\delta^*_1 = 1, T_1 > T_2] - \frac{1}{2}.$ 

Consider the kernel

$$\phi_{4}(T_{i}, \delta_{i}, T_{j}, \delta_{j}) = \begin{cases} 1 & \text{if } \delta^{*}_{i} = 1, T_{i} > T_{j} \\ & \text{or } \delta^{*}_{j} = 1, T_{i} < T_{j} \\ 0 & \text{otherwise} \end{cases}$$

The corresponding U-statistic is given by

$$U_4 = \frac{1}{\binom{n}{2}} \sum_{1 \le i < j \le n} \phi_4(T_i, \delta_i, T_j, \delta_j).$$

$$\binom{n}{2}U_4 = \sum_{i=1}^n (R_i - 1)\delta^*_i.$$

This statistic was earlier proposed by Bagai, Deshpande and Kochar (1989) to test for equality of failure rates of independent latent competing risks.

Aly , Kochar and Mckeague (1994) proposed Kolmogrov-Smirnov type tests for testing the equality of two competing risks against the alternatives  $H_1$  and  $H_2$ .

$$\psi^*(t) = F(2,t) - F(1,t)$$
  
= 
$$\int_0^t S(u)(\lambda(2,u) - \lambda(1,u)) du.$$

Under  $H_0, \ \psi^*(t) = 0.$ 

 $H_1$  holds iff  $\psi^*(t)$  is nondecreasing in t .

Let  $\psi_n^*(t)$  be its empirical estimator.

$$D_{1n} = \sup_{0 \le s < t < \infty} \{ \psi_n^*(t) - \psi_n^*(s) \}.$$

Large values of the statistic are significant.

Exact null distribution of  $D_{1n}$  is given below for t = 1, ..., n + 1,

 $P[nD_{1n} < t]$ =  $\frac{2}{2t+1} \sum_{j=0}^{2t} (\cos \frac{j\pi}{2t+1}) (\sin \frac{j\pi(t+1)}{2t+1})$ ×  $(1 + \cos \frac{j\pi}{2t+1}) (\frac{1-(-1)^j}{2}) / \sin \frac{j\pi}{2t+1}.$ 

Asymptotic null distribution of  $D_{1n}$  is given by

$$\sqrt{n}D_{1n} \xrightarrow{\mathcal{L}} \sup_{0 \le x \le 1} |W(x)|,$$

where  $\{W(t), t \ge 0\}$  is a standard Brownian motion.

The asymptotic 0.90, 0.95, 0.99 quantiles of  $\sqrt{n}D_{1n}$  were found to be 1.96, 2.241, 2, 807, respectively.

For a general two sided alternative  $F(1,t) \neq F(2,t)$  for some t or equivalently  $\lambda_1(t) \neq \lambda_2(t)$  for some t, then one can use the Kolmogrov-Smirnov type statistic

 $D_n = \sup_{t \ge 0} |\psi_n^*(t)|.$ 

Under 
$$H_0, \sqrt{n}D_n \xrightarrow{\mathcal{L}} \sup_{0 \le x \le 1} |W(x)|$$
.

This test is consistent against arbitrary departures from  $H_0$ .

For testing  $H_0$  against  $H_2$ , Aly , Kochar and McKeague (1994) proposed the statistic

 $D_{2n} = \sup_{0 \le t\infty} \psi_n(t).$ 

Large values of  $D_{2n}$  are significant for testing  $H_0$  against  $H_2$ .

The exact nulltribution of  $D_{2n}$  is

$$P[nD_{2n} = k] = \frac{1}{2^n} {\binom{n}{\left[\frac{n-k}{2}\right]}}, \quad k = 0, 1, \dots, n.$$

The asymptotic null distribution is

$$P[\sqrt{n}D_{2n} > x] \rightarrow 2(1 - \Phi(x)), \ x \ge 0,$$

where  $\Phi$  is the standard normal distribution function.

#### Remarks

(i) The various tests are consistent against their intended alternatives.

(ii) The tests are distribution-free under  $H_0$ . The null distribution of the tests  $U_3$  and  $U_4$  is same as in the case of independent latent failures (see, Deshpande and Dewan (2003)).

(iii) We can also use these tests for the hypothesis  $\lambda_1(t) = \lambda_2(t)$  against the alternative that cause-specific hazards are ordered.

(iv) T and  $\delta$  continue to be independent under the null hypothesis of bivariate symmetry .

(v) The statistic  $U_2$  puts more weight on the middle observations and is less sensitive to the observations in the beginning and the end of the experiment.

 $U_3$  puts more weight to later observations and

 $U_4$  puts higher weight to observations in the beginning.

(vi) The statistics  $U_2, U_3, U_4$  are all linear combinations of the sign statistic and the Wilcoxon-signed rank type statistic.

(vii) Tests proposed by Aly, Kochar and Mckeague (1994) can be extended to the case of multiple risks in which any two of the causespecific risks are to be compared. The statistic can be modified to test dominance of one risk over the other in a specified interval.

Censored data

Let C be the censoring random variable independent of the latent failure times X and Y.

The survival function of C is  $S_c$  and assume that  $S_C(t) > 0$  for all t.

The available information consists of  $(\tilde{T}_i, \tilde{\delta}_i), i = 1, 2, \cdots, n$ 

 $\tilde{T} = \min(T, C)$  and  $\tilde{\delta} = \delta^* I(T \leq C)$ .

Aly et al (1994) generalised the function  $\psi^*$  so as to capture departures of  $H_0$  from  $H_2$  in case of censored data.

$$\phi(t) = \int_0^t S(u-)(S_C(u-))^{\frac{1}{2}}(\lambda(2,u) - \lambda(1,u))du$$

which is the  $\psi^*$  function when there is no censoring.

The integrand  $S_C(u-)^{1/2}$  is the function required to compensate for censoring in order that the *D* statistics remain asymptotically distribution free.

Under  $H_0, \phi(t) = 0$ .

 $H_1$  holds iff  $\phi(t)$  is increasing in t.

The relevant statistic is

$$D_{3n} = \sup_{0 \le s < t < \infty} \{\phi_n(t) - \phi_n(s)\}.$$

An obvious choice of  $\phi_n$  is

$$\phi_n(t) = \int_0^t \hat{S}(u-) (\hat{S}_C(u-))^{1/2} d(\hat{\Lambda}_1 - \hat{\Lambda}_2)(u)$$
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 $\hat{S}$  and  $\hat{S}_{C}$  are the product limit estimators of  $\bar{S}$  and  $S_{C}\text{,}$ 

 $\widehat{\Lambda}_j$  is the Aalen estimator of the cumulative CSHR function  $\Lambda_j(t) = \int_0^t \lambda(j, u) \, du$ .

$$\hat{\Lambda}(j,t) = \sum_{i \mid \tilde{T}_i \leq t} \frac{I(\tilde{\delta}_i = j)}{R_i}$$

where  $R_i = \#\{k : \tilde{T}_k \ge \tilde{T}_i\}$  is the risk set at time  $\tilde{T}_i$ .

Large values of the statistic are significant.

Since  $\phi(t) > 0$  for some t under  $H_2$ , a suitable test procedure is based on large values of

 $D_{4n} = \sup_{0 < t < \infty} \phi_n(t).$ 

Aly et al (1994) showed that  $D_{3n}$  and  $D_{4n}$  are asymptotically distribution-free with the same limiting distributions as those obtained in the uncensored case.

A suitable modification of  $U_2$  to censored data is given by the statistic

$$K_n = \int_{0 < x < y < \infty} (\phi_n(x) - \phi_n(y)) \, d\widehat{S}(x) \, d\widehat{S}(y)$$

where  $\phi_n$  and  $\hat{S}$  are as defined above.

 $K_n$  can be expressed as

$$K_n = \sum_{i \mid \tilde{\delta}_i \neq 0} \phi_n(i) (2\hat{S}(\tilde{T}_{(i-1)}) - 1) (\hat{S}(\tilde{T}_{(i-1)}) - 1) (\hat{S}(\tilde{T}_$$

Large values of  $K_n$  are significant for testing  $H_0$  against  $H_1$ .

Aly et al (1994) showed that, under  $H_0$ ,

 $n^{1/2}\phi_n \xrightarrow{\mathcal{L}} W(S(.))$ 

where W(.) is the standard Brownian Motion.

Using the continuous mapping theorem, we have

$$n^{1/2}K_n \xrightarrow{\mathcal{L}} N(0,\sigma^2)$$

 $\sigma^2 = \frac{1}{30}$ 

Sun and Tiwari (1995) modified the statistic  $U_3$  so that it can be used for censored data.

$$V = \int_0^\infty (F(2,t) - F(1,t)) dH(t) = \int_0^\infty [S(t)]^2 d(\Lambda_2(t) - \Lambda_1(t))$$

A natural estimator of V is given by  $V_n$  where

$$V_n = \int_0^\infty [\widehat{S}(t-)]^2 d(\widehat{\Lambda}_2(t) - \widehat{\Lambda}_1(t))$$

where  $\widehat{S}(t)$  and  $\widehat{\Lambda}_j$  are as defined above.

In the absence of censoring  $V_n$  reduces to the statistic  $U_3$ .

Then Sun and Tiwari (1995) proved the following theorem.

**Theorem:** Let  $K(t) = 1 - S(t)S_C(t)$ ,  $\tau_K = \sup\{t : K(t) < 1\}$ . If  $\int_0^{\tau_K} \frac{dH(t)}{S_C(t)} < \infty$ , then

$$\sqrt{n}(V_n-V) \xrightarrow{\mathcal{L}} N(0,\sigma^2) \ asn \to \infty.$$

Under  $H_0$ ,

$$\sigma^2 = \int_0^\infty \frac{S^2(t)}{S_C(t)} dH(t).$$

A consistent estimator  $\hat{\sigma}^2$  of  $\sigma^2$  is obtained by replacing S ans  $S_C$  by their consistent Kaplan Meier-estimators.

When there is no censoring  $\sigma^2 = \frac{1}{3}$ .

The test rejects for large values of the statistic.

The approximate power of this test of size  $\alpha$  is equal to  $1 - \Phi(z_{\alpha} - \sqrt{n}V/\hat{\sigma})$ , where  $z_{\alpha}$  is the upper  $\alpha$  percentile of standard normal distribution.

### SIMULATION RESULTS

Random samples were generated from absolutely continuous bivariate exponential (ACBVE) due to Block and Basu (1974) with density

$$f(x,y) = \begin{cases} \frac{\lambda\lambda_1(\lambda_2+\lambda_0)}{\lambda_1+\lambda_2}e^{-\lambda_1x-(\lambda_2+\lambda_0)y} & \text{if } x < y, \\ \frac{\lambda\lambda_2(\lambda_1+\lambda_0)}{\lambda_1+\lambda_2}e^{-\lambda_2y-(\lambda_1+\lambda_0)x} & \text{if } x > y, \end{cases}$$

where  $(\lambda_0, \lambda_1, \lambda_2)$  are the parameters and  $\lambda = \lambda_0 + \lambda_1 + \lambda_2$ . The CSHR's are

$$\lambda_j(t) = \frac{\lambda_j \lambda}{\lambda_1 + \lambda_2}, \quad j = 1, 2.$$

Under  $H_1 \quad \lambda_1 < \lambda_2$ .

X and Y are independent if and only if  $\lambda_0 = 0$ .

We set  $\lambda_1 = 1$  and consider  $\lambda_2 = 1.0, 1.4, 1.8, 2.2$ indicating larger and larger departures from  $H_0$ .

The case  $\lambda_2 = 1.0$  corresponds to the null hypothesis. n = 100 and there are 10000 replications.

	$\lambda_2$			
Test	1.0	1.4	1.8	2.2
$D_1$	3.76	41.98	82.53	96.83
D <sub>2</sub>	4.85	47.71	86.98	98.14
U2	5.09	44.60	83.96	96.92
U3	4.79	43.06	80.54	95.42
U4	4.99	42.67	80.81	95.77
Sign	4.39	49.50	88.29	98.66

Next we look at the censored case. The censoring distribution was exponential with parameters 1 and 3, respectively. We use asymptotic critical levels of 5 percent. Results are based on 5,000 replications.

Observed levels and powers of  $K_n$  at an asymptotic level of 5 percent. The underlying distribution of (X, Y) is Block and Basu (1974) ACBVE with  $\lambda_1 = 1$ .

## (a) CENSORED (EXP(1))

	n=50		n=100	
$\lambda_2$	$\lambda_0 = 0$	$\lambda_0 = 1$	$\lambda_0 = 0$	$\lambda_0 = 1$
1.0	.0218	.0312	.0360	.0376
1.5	.1864	.2192	.3732	.4302
2.0	.4482	.5080	.7862	.8342
2.5	.6928	.7414	.9546	.9704

# (b) CENSORED (EXP(3))

	n=50		n=100	
$\lambda_2$	$\lambda_0 = 0$	$\lambda_0 = 1$	$\lambda_0 = 0$	$\lambda_0 = 1$
1.0	.0048	.0124	.0084	.0172
1.5	.0342	.0762	.1012	.1834
2.0	.1202	.1986	.3216	.4860
2.5	.2496	.3774	.5862	.7344

From the table it is clear that the asymptotic critical levels give conservative tests for the censored case, with the effect increasing as the censoring becomes more severe.

There is slight effect on the levels or the power due to lack of independence of X and Y in the presence of censoring.

The results are comparable with the test proposed by Aly et al (1994) for the lightly censored case.

Test for independence of T and  $\delta$ 

The nature of dependence between T and  $\delta$  is crucial and useful in modelling competing risks data via sub-distribution/subsurvival functions.

If T and  $\delta$  are independent then  $S_i(t) = pr(\delta = i)S(t)$ , allowing the study of the failure times and the causes (risks) of failure separately.

The hypothesis of equality of incidence functions or that of cause-specific hazard rates reduces to testing whether  $pr(\delta = 1) = pr(\delta = 0) = 1/2$ .

This simplifies the study of competing risks to a great extent.

Dewan, Deshpande and Kulathinal (2004) studied the properties of the conditional probability functions

 $\Phi_i(t) = pr(\delta = i \mid T \ge t) = S_i(t)/S(t), \ i = 1, 2$  $\Phi_i^*(t) = pr(\delta = i \mid T < t) = F_i(t)/H(t), \ i = 1, 2$ 1, 2. (i) T and  $\delta$  are independent iff  $\Phi_1(t) = P[\delta = 1]$  or  $\Phi_2^*(t) = 1 - P[\delta = 1]$ 

(ii) T and  $\delta$  are PQD iff  $\Phi_1(t) \ge P[\delta = 1]$  or  $\Phi_2^*(t) \ge 1 - P[\delta = 1]$ 

(iii)  $\delta$  is Right Tail Increasing in T iff  $\Phi_1(t)$  is increasing in t.

(iv)  $\delta$  is Left Tail Decreasing in T iff  $\Phi_2^*(t)$  is decreasing in t.

They considered the problem of testing  $H_0$ : T and  $\delta$  are independent which is equivalent to

 $H_0: \Phi_1(t)$  is a constant

against various alternative hypotheses which characterise the properties of  $\Phi_1(t)$  and  $\Phi_0^*(t)$ :

 $H_1: \Phi_1(t)$  is not a constant

 $H_2: \Phi_1(t) \ge P[\delta = 1]$  for all t with strict inequality f

 $H_3: \Phi_1(t)$  is a monotone nondecreasing function of

 $H_4: \Phi_0^*(t)$  is a monotone nonincreasing function of

If the hypothesis of independence is accepted then one can simplify the model and study the failure time and cause of failure separately.

If the hypothesis is rejected then a suitable model under specific dependence between T and  $\delta$  in terms of the incidence functions is needed.

The tests constructed for the two risk case cannot be straightway extended to the case of more than 2 causes of failure.

For example, in the most commonly cited mortality data given in Hoel (1972), the data were obtained from a laboratory experiment on two groups of RFM strain male mice which had received a radiation dose of 300 r at an age of 5-6 weeks.

The first group of mice lived in a conventional laboratory environment, while the second group was in a germ-free environment. The causes of death were grouped into three classes - thymic lymphoma (risk=1), reticulum cell sarcoma (risk=2) and all other causes (risk=3).

Suppose we have information on  $(T, \delta)$ , where  $\delta$  takes three values 1, 2, 3.

Let 
$$\phi_i(t) = \frac{S_i(t)}{S(t)}$$
.

 $(T,\delta)$  are PQD iff

$$\phi_3(t) \ge \phi_3(0) = \phi_3,$$

and

 $\phi_1(t) \le \phi_1(0) = \phi_1,$ 

Similarly  $\delta$  is RTI in T iff

 $\phi_3(t)$  is increasing in t

and

 $\phi_1(t)$  is decreasing in t.

#### **Tests For PQD**

$$\int_0^\infty [S_3(t) - \phi_3 S(t)] dF(t) = P[T_2 > T_1, \delta_2 = 3]$$
  
and

$$\int_0^\infty [S_1(t) - \phi_1 S(t)] dF(t) = P[T_2 > T_1, \delta_2 = 1].$$

Consider

$$\psi_{1}(T_{i}, \delta_{i}, T_{j}, \delta_{j}) = \begin{cases} 1 & \text{if } T_{j} > T_{i}, \delta_{i} = 1, \delta_{j} = 3, \\ T_{j} > T_{i}, \delta_{i} = 2, \delta_{j} = 3, \\ T_{j} > T_{i}, \delta_{i} = 3, \delta_{j} = 3, \\ \text{or if } T_{i} > T_{j}, \delta_{i} = 3, \delta_{j} = 1, \\ T_{i} > T_{j}, \delta_{i} = 3, \delta_{j} = 2, \\ T_{i} > T_{j}, \delta_{i} = 3, \delta_{j} = 3, \\ 0 & \text{otherwise.} \end{cases}$$

Let  $U_1$  be the corresponding U-statistic.

Further consider

$$\psi_{2}(T_{i},\delta_{i},T_{j},\delta_{j}) = \begin{cases} 1 & \text{if } T_{j} > T_{i},\delta_{i} = 1, \delta_{j} = 1, \\ T_{j} > T_{i},\delta_{i} = 2,\delta_{j} = 1, \\ T_{j} > T_{i},\delta_{i} = 3,\delta_{j} = 1, \\ \text{or if } T_{i} > T_{j},\delta_{i} = 1,\delta_{j} = 1, \\ T_{i} > T_{j},\delta_{i} = 1,\delta_{j} = 2, \\ T_{i} > T_{j},\delta_{i} = 1,\delta_{j} = 3, \\ 0 & \text{otherwise.} \end{cases}$$

Let  $U_2$  be the corresponding U-statistic. Then  $U_1 - U_2$  can be used to test for PQD. Large values of the statistic are significant.  $E(U_1) = 2 \int_0^\infty S_3(t) dF(t), \quad E(U_2) = 2 \int_0^\infty S_1(t) dF(t).$ In particular under the null hypotheses  $E(U_1) = \phi_3, \ E(U_2) = \phi_1.$ Under  $H_0$ ,  $\sigma_1^2 = \frac{1}{3}\phi_3(1-\phi_3),$  $\sigma_2^2 = \frac{1}{3}\phi_1(1-\phi_1).$ Theorem :  $\sqrt{n}(U_i - E(U_i)) \xrightarrow{\mathcal{L}} N(0, \sigma_i^2), i =$ 1,2 as  $n \rightarrow \infty, i = 1, 2$ . Under  $H_0$ 

$$Cov(U_1, U_2) = -\frac{2\phi_1\phi_3}{3}\frac{2n-1}{n(n-1)}.$$

Hence , under  $H_0$  , limiting variance of  $\sqrt{n}(U_1-U_2)$  is given by

 $\sigma^2 = \frac{4}{3}(\phi_1 + \phi_3) - \frac{4}{3}(\phi_1 - \phi_3)^2.$ 

And  $\sqrt{n}(U_1 - U_2) \xrightarrow{\mathcal{L}} N(0, \sigma^2)$ , as  $n \to \infty$ .

One can use asymptotic critical points for testing purposes.

One can easily find the exact distribution of  $U_1, U_2$ .

Even the tests for independence of T and  $\delta$  are linear combinations of the sign statistic and the signed rank statistic .

MODELS TO FIT THIS SET UP?