NAG Library Routine Document

G12ABF

Note: before using this routine, please read the Users' Note for your implementation to check the interpretation of **bold italicised** terms and other implementation-dependent details.

1 Purpose

G12ABF calculates the rank statistics, which can include the logrank test, for comparing survival curves.

2 Specification

3 Description

A survivor function, S(t), is the probability of surviving to at least time t. Given a series of n failure or right-censored times from g groups G12ABF calculates a rank statistic for testing the null hypothesis

$$H_0: S_1(t) = S_2(t) = \cdots = S_q(t), t \le \tau$$

where τ is the largest observed time, against the alternative hypothesis

 H_1 : at least one of the $S_i(t)$ differ, for some $t \leq \tau$.

Let t_i , for $i = 1, 2, ..., n_d$, denote the list of distinct failure times across all g groups and w_i a series of n_d weights. Let d_{ij} denote the number of failures at time t_i in group j and n_{ij} denote the number of observations in the group j that are known to have not failed prior to time t_i , i.e., the size of the risk set for group j at time t_i . If a censored observation occurs at time t_i then that observation is treated as if the censoring had occurred slightly after t_i and therefore the observation is counted as being part of the risk set at time t_i . Finally let

$$d_i = \sum_{j=1}^g d_{ij}$$
 and $n_i = \sum_{j=1}^g n_{ij}$.

The (weighted) number of observed failures in the jth group, O_i , is therefore given by

$$O_j = \sum_{i=1}^{n_d} w_i d_{ij}$$

and the (weighted) number of expected failures in the jth group, E_j , by

$$E_j = \sum_{i=1}^{n_d} w_i \frac{n_{ij} d_i}{n_i}.$$

If x denotes the vector of differences $x = (O_1 - E_1, O_2 - E_2, \dots, O_q - E_q)$ and

$$V_{jk} = \sum_{i=1}^{n_d} w_i^2 \left(\frac{d_i(n_i - d_i) (n_i n_{ik} I_{jk} - n_{ij} n_{ik})}{n_i^2 (n_i - 1)} \right)$$

where $I_{jk} = 1$ if j = k and 0 otherwise, then the rank statistic, T, is calculated as

$$T = rV^-r^T$$

G12ABF NAG Library Manual

where V^- denotes a generalized inverse of the matrix V. Under the null hypothesis, $T \sim \chi^2_{\nu}$ where the degrees of freedom, ν , is taken as the rank of the matrix V.

4 References

Gross A J and Clark V A (1975) Survival Distributions: Reliability Applications in the Biomedical Sciences Wiley

Kalbfleisch J D and Prentice R L (1980) The Statistical Analysis of Failure Time Data Wiley

Rostomily R C, Duong D, McCormick K, Bland M and Berger M S (1994) Multimodality management of recurrent adult malignant gliomas: results of a phase II multiagent chemotherapy study and analysis of cytoreductive surgery Neurosurgery 35 378

5 **Arguments**

N - INTEGER 1: Input

On entry: n, the number of failure and censored times.

Constraint: $N \ge 2$.

2: T(N) – REAL (KIND=nag_wp) array Input

On entry: the observed failure and censored times; these need not be ordered.

Constraint: $T(i) \neq T(j)$ for at least one $i \neq j$, for i = 1, 2, ..., N and j = 1, 2, ..., N.

3: IC(N) – INTEGER array Input

On entry: IC(i) contains the censoring code of the ith observation, for i = 1, 2, ..., N.

the ith observation is a failure time.

IC(i) = 1

the *i*th observation is right-censored.

Constraints:

$$IC(i) = 0$$
 or 1, for $i = 1, 2, ..., N$;
 $IC(i) = 0$ for at least one i .

GRP(N) - INTEGER array 4:

Input

Input

On entry: GRP(i) contains a flag indicating which group the ith observation belongs in, for $i = 1, 2, \dots, N$.

Constraints:

```
1 \leq GRP(i) \leq NGRP, for i = 1, 2, ..., N;
each group must have at least one observation.
```

NGRP - INTEGER 5:

On entry: g, the number of groups.

Constraint: $2 \le NGRP \le N$.

FREQ - CHARACTER(1)

Input

On entry: indicates whether frequencies are provided for each time point.

FREO = 'F'

Frequencies are provided for each failure and censored time.

G12ABF.2 Mark 26 FREQ = 'S'

The failure and censored times are considered as single observations, i.e., a frequency of 1 is assumed.

Constraint: FREQ = 'F' or 'S'.

7: IFREQ(*) – INTEGER array

Input

Note: the dimension of the array IFREQ must be at least N if FREQ = 'F'.

On entry: if FREQ = 'F', IFREQ(i) must contain the frequency (number of observations) to which each entry in T corresponds.

If FREQ = 'S', each entry in T is assumed to correspond to a single observation, i.e., a frequency of 1 is assumed, and IFREQ is not referenced.

Constraint: if FREQ = 'F', IFREQ(i) ≥ 0 , for i = 1, 2, ..., N.

8: WEIGHT - CHARACTER(1)

Input

On entry: indicates if weights are to be used.

WEIGHT = 'U'

All weights are assumed to be 1.

WEIGHT = 'W'

The weights, w_i are supplied in WT.

Constraint: WEIGHT = 'U' or 'W'.

9: WT(*) - REAL (KIND=nag_wp) array

Input

Note: the dimension of the array WT must be at least LDN if WEIGHT = 'W'.

On entry: if WEIGHT = 'W', WT must contain the n_d weights, w_i , where n_d is the number of distinct failure times.

If WEIGHT = 'U', WT is not referenced and $w_i = 1$ for all i.

Constraint: if WEIGHT = 'W', WT(i) \geq 0.0, for $i = 1, 2, ..., n_d$.

10: TS - REAL (KIND=nag_wp)

Output

On exit: T, the test statistic.

11: DF - INTEGER

Output

On exit: ν , the degrees of freedom.

12: P - REAL (KIND=nag wp)

Output

On exit: $P(X \ge T)$, when $X \sim \chi^2_{\nu}$, i.e., the probability associated with TS.

13: OBSD(NGRP) – REAL (KIND=nag wp) array

Output

On exit: O_i , the observed number of failures in each group.

14: EXPT(NGRP) - REAL (KIND=nag wp) array

Output

On exit: E_i , the expected number of failures in each group.

15: ND - INTEGER

Output

On exit: n_d , the number of distinct failure times.

G12ABF NAG Library Manual

16: DI(LDN) – INTEGER array

Output

On exit: the first ND elements of DI contain d_i , the number of failures, across all groups, at time t_i .

17: NI(LDN) – INTEGER array

Output

On exit: the first ND elements of NI contain n_i , the size of the risk set, across all groups, at time t_i .

18: LDN – INTEGER

Input

On entry: the size of arrays DI and NI. As $n_d \le n$, if n_d is not known a priori then a value of N can safely be used for LDN.

Constraint: LDN $\geq n_d$, the number of unique failure times.

19: IFAIL – INTEGER

Input/Output

On entry: IFAIL must be set to 0, -1 or 1. If you are unfamiliar with this argument you should refer to Section 3.4 in How to Use the NAG Library and its Documentation for details.

For environments where it might be inappropriate to halt program execution when an error is detected, the value -1 or 1 is recommended. If the output of error messages is undesirable, then the value 1 is recommended. Otherwise, if you are not familiar with this argument, the recommended value is 0. When the value -1 or 1 is used it is essential to test the value of IFAIL on exit.

On exit: IFAIL = 0 unless the routine detects an error or a warning has been flagged (see Section 6).

6 Error Indicators and Warnings

If on entry IFAIL = 0 or -1, explanatory error messages are output on the current error message unit (as defined by X04AAF).

Errors or warnings detected by the routine:

```
IFAIL = 1
```

On entry, $N = \langle value \rangle$. Constraint: $N \ge 2$.

IFAIL = 2

On entry, all the times in T are the same.

IFAIL = 3

On entry, $IC(\langle value \rangle) = \langle value \rangle$. Constraint: IC(i) = 0 or 1.

IFAIL = 4

On entry, $GRP(\langle value \rangle) = \langle value \rangle$ and $NGRP = \langle value \rangle$. Constraint: $1 \leq GRP(i) \leq NGRP$.

IFAIL = 5

On entry, NGRP = $\langle value \rangle$ and N = $\langle value \rangle$. Constraint: 2 < NGRP < N.

IFAIL = 6

On entry, FREQ had an illegal value.

G12ABF.4 Mark 26

```
IFAIL = 7
```

On entry, IFREQ($\langle value \rangle$) = $\langle value \rangle$. Constraint: IFREQ(i) \geq 0.

IFAIL = 8

On entry, WEIGHT had an illegal value.

IFAIL = 9

On entry, $WT(\langle value \rangle) = \langle value \rangle$. Constraint: $WT(i) \ge 0.0$.

IFAIL = 11

The degrees of freedom are zero.

IFAIL = 18

On entry, LDN = $\langle value \rangle$. Constraint: LDN $\geq \langle value \rangle$.

IFAIL = 31

On entry, all observations are censored.

IFAIL = 41

On entry, group $\langle value \rangle$ has no observations.

IFAIL = -99

An unexpected error has been triggered by this routine. Please contact NAG.

See Section 3.9 in How to Use the NAG Library and its Documentation for further information.

IFAIL = -399

Your licence key may have expired or may not have been installed correctly.

See Section 3.8 in How to Use the NAG Library and its Documentation for further information.

IFAIL = -999

Dynamic memory allocation failed.

See Section 3.7 in How to Use the NAG Library and its Documentation for further information.

7 Accuracy

Not applicable.

8 Parallelism and Performance

G12ABF is threaded by NAG for parallel execution in multithreaded implementations of the NAG Library.

G12ABF makes calls to BLAS and/or LAPACK routines, which may be threaded within the vendor library used by this implementation. Consult the documentation for the vendor library for further information.

Please consult the X06 Chapter Introduction for information on how to control and interrogate the OpenMP environment used within this routine. Please also consult the Users' Note for your implementation for any additional implementation-specific information.

G12ABF NAG Library Manual

9 Further Comments

The use of different weights in the formula given in Section 3 leads to different rank statistics being calculated. The logrank test has $w_i=1$, for all i, which is the equivalent of calling G12ABF when WEIGHT = 'U' . Other rank statistics include Wilcoxon ($w_i=n_i$), Tarone-Ware ($w_i=\sqrt{n_i}$) and Peto-Peto ($w_i=\tilde{S}(t_i)$ where $\tilde{S}(t_i)=\prod\limits_{t_j\leq t_i}\frac{n_j-d_j+1}{n_j+1}$) amongst others.

Calculation of any test, other than the logrank test, will probably require G12ABF to be called twice, once to calculate the values of n_i and d_i to facilitate in the computation of the required weights, and once to calculate the test statistic itself.

10 Example

This example compares the time to death for 51 adults with two different types of recurrent gliomas (brain tumour), astrocytoma and glioblastoma, using a logrank test. For further details on the data see Rostomily *et al.* (1994).

10.1 Program Text

```
Program g12abfe
     G12ABF Example Program Text
1
!
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!
      .. Use Statements ..
     Use nag_library, Only: g12abf, nag_wp
!
      .. Implicit None Statement ..
     Implicit None
      .. Parameters ..
!
                                       :: nin = 5, nout = 6
     Integer, Parameter
      .. Local Scalars ..
!
     Real (Kind=nag_wp)
                                        :: p, ts
                                        :: df, i, ifail, ldn, lfreq, lwt, n, nd, ngrp
     Integer
     Character (1)
                                        :: freq, weight
!
      .. Local Arrays ..
     Real (Kind=nag_wp), Allocatable :: expt(:), obsd(:), t(:), wt(:)
     Integer, Allocatable
                                       :: di(:), grp(:), ic(:), ifreq(:),
                                          ni(:)
      .. Executable Statements ..
!
     Write (nout,*) 'G12ABF Example Program Results'
     Write (nout,*)
     Skip heading in data file
     Read (nin,*)
     Read in the problem size
!
     Read (nin,*) n, ngrp, freq, weight
     If (freq=='F' .Or. freq=='f') Then
        lfreq = n
     Else
       lfreq = 0
      If (weight=='W' .Or. weight=='w') Then
       lwt = n
     Else
       lwt = 0
     End If
     Allocate (t(n),ic(n),grp(n),ifreq(lfreq),obsd(ngrp),expt(ngrp),di(ldn), &
        ni(ldn),wt(lwt))
     Read in data
     If (lfreq==0) Then
```

G12ABF.6 Mark 26

```
Read (nin,*)(t(i),ic(i),grp(i),i=1,n)
        Read (nin,*)(t(i),ic(i),grp(i),ifreq(i),i=1,n)
      End If
      Calculate the statistic
      ifail = 0
      Call g12abf(n,t,ic,grp,ngrp,freq,ifreq,weight,wt,ts,df,p,obsd,expt,nd,
        di,ni,ldn,ifail)
!
     Display results
      Write (nout, 99999) 'Observed', 'Expected'
      Write (nout,99998)('Group',i,obsd(i),expt(i),i=1,ngrp)
      Write (nout,*)
      Write (nout, 99996) 'No. Unique Failure Times = ', nd
      Write (nout,*)
                                                    = ', ts
= ', df
= ', p
      Write (nout,99997) 'Test Statistic
      Write (nout, 99996) 'Degrees of Freedom
      Write (nout, 99997) 'p-value
99999 Format (11X,A,2X,A)
99998 Format (1X,A,1X,I1,1X,F8.2,2X,F8.2)
99997 Format (1X,A,1X,F8.4)
99996 Format (1X,A,1X,I3)
    End Program g12abfe
```

10.2 Program Data

```
G12ABF Example Program Data
51 2 'S' 'U' :: N, NGRP, FREQ, WEIGHT
 6.0 0 1
13.0 0
        1
21.0 0
         1
30.0 0
         1
31.0 1
         1
37.0 0
38.0 0
         1
47.0
      1
         1
49.0 0
         1
50.0 0
         1
63.0 0
         1
79.0 0
         1
80.0
      1
         1
82.0 1
         1
82.0 1
         1
86.0 0
         1
98.0
      0
149.0 1
         1
202.0 0
         1
219.0 0
         1
10.0
      0
         2
10.0 0
         2
12.0 0
         2
13.0 0
         2
14.0 0
         2
15.0
      0
         2
         2
16.0 0
17.0 0
18.0 0
         2
20.0
      0
         2
24.0 0
         2
24.0 0
         2
25.0 0
         2
28.0
     0
         2
30.0
     0
         2
33.0
      0
         2
34.0 1
35.0 0
        2
37.0 0
40.0 0 2
```

G12ABF NAG Library Manual

```
40.0 0 2

40.0 1 2

46.0 0 2

48.0 0 2

70.0 1 2

76.0 0 2

81.0 0 2

82.0 0 2

91.0 0 2

112.0 0 2

181.0 0 2 :: T,IC,GRP
```

10.3 Program Results

G12ABF Example Program Results

```
Observed Expected
Group 1
          14.00
                   22.48
                    19.52
          28.00
Group 2
No. Unique Failure Times =
                             36
Test Statistic
                              7.4966
Degrees of Freedom
                        =
                              1
p-value
                              0.0062
```

G12ABF.8 (last)

Mark 26