Statistical methodologies for goodness-of-fit: a comparative analysis of three established approaches

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**Goodness-of-fit tests** can be applied: available for both complete and censored data.

- Although non-parametric approaches are commonly used for this kind of data, parametric approaches also play an important role in survival analysis.
- Adaptations of classic goodness-of-fit tests to censored data are available.

Data

distribution

Statistical models and procedures

# <span id="page-4-0"></span>Tests based on the empirical distribution function

Test the hypothesis:

 $H_0$ :  $F(t) = F_0(t)$  $H_1$  :  $F(t) \neq F_0(t)$ , for all  $t \geq 0$ 

Most common ones:

- Kolmogorov-Smirnov
- Cramér-von Mises
- Anderson-Darling

When applying these tests to censored data, the empirical distribution function is replaced by an appropriate estimate of the distribution function.

<span id="page-5-0"></span>Adaptation proposed by Fleming et al. given by:

$$
D_n = \sup_t |\hat{F}_n(t) - F_0(t)|
$$

Where:

- $\hat{\mathcal{F}}_n(t)$  is the estimation of the empirical distribution function of the data.
- $\bullet$  *n* is the sample size.

For **right-censored data**  $\hat{\mathsf{F}}_n$ , is replaced by  $\hat{\mathsf{F}}_n=1-\hat{\mathsf{S}}_n=1-\mathrm{e}^{-\hat{\Lambda}_n},$  where  $\hat{\Lambda}_n$  denotes the Nelson-Aalen estimator of the cumulative hazard function.

# <span id="page-6-0"></span>Kolmogorov-Smirnov

As a result:

$$
\hat{D}_n = \sup_{0 \leq t \leq t_m} |\hat{F}_n(t) - F_0(t)| = \sup_{0 \leq t \leq t_m} \left| \int_0^t \frac{\hat{S}_n(t) S_0(s)}{\hat{S}_n(s)} d\left[\hat{\Lambda}_n(s) - \Lambda_0(s)\right] \right|,
$$

where  $S_0$  and  $\Lambda_0$  are, respectively, the survival and the cumulative hazard function of the hypothesized distribution and  $t_m$  is the largest observed time in the sample.

## <span id="page-7-0"></span>Cramér-von Mises

The Cramér-von Mises statistic is given by:

$$
M_n = n \int_{-\infty}^{+\infty} \left(\hat{F}_n(t) - F_0(t)\right)^2 dF_0(t),
$$

where  $\hat{F}_n$  is:

- The empirical distribution function when data is complete.
- $\bullet$  1 minus the Kaplan-Meier estimator of the survival function  $(\hat{\digamma}_n = 1 \hat{S}_n)$  if the data has random right-censorship.

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Cramér-von Mises

When data has right-censored observations:

$$
\hat{M}_n = n_r \sum_{j=1}^{n_r+1} \hat{F}_n(u_{(j-1)})(u_{(j)} - u_{(j-1)}) \left( \hat{F}_n(u_{(j-1)}) - (u_{(j)} + u_{(j-1)}) \right) + \frac{n_r}{3},
$$

where,  $Y_1,\ldots,Y_{n_r}$  are the  $n_r$  observed failure times and  $u_{(i)}\ =\ F_0(Y_{(i)})$  if we transform the order statistic  $\mathsf{Y}_{(1)},\ldots,\mathsf{Y}_{(n_r)}$  into  $\mathsf{Uniform}(0,1)$  random variables  $u_{(1)}, \ldots, u_{(n_r)}.$ 

The asymptotic distribution of  $\hat{M}_n$  is not easily implemented.

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# Anderson-Darling

Anderson-Darling statistics:

$$
A_n = n \int_{-\infty}^{+\infty} (\hat{F}_n(t) - F_0(t))^2 \frac{dF_0(t)}{F_0(t)(1 - F_0(t))}.
$$

where:

- $\hat{\mathsf{F}}_n$  is the empirical distribution function when data are complete.
- $\hat{\mathsf{F}}_n$  is 1 minus the Kaplan-Meier estimator of the survival function  $(\hat{\mathcal{F}}_n = 1 - \hat{\mathcal{S}}_n)$  if the data has random right-censorship.

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### Anderson-Darling

When data has right-censored observations:

$$
\hat{A}_n = -n_r + n_r \sum_{j=1}^{n_r} (\hat{F}_n(u_{(j-1)}) - 1)^2 \left[ \log |1 - u_{(j-1)}| - \log |1 - u_{(j)}| \right] + n_r \sum_{j=1}^{n_r - 1} \hat{F}_n^2(u_{(j)}) \left[ \log |u_{(j+1)}| - \log |u_{(j)}| \right] - n_r \log |u_{(n)}|.
$$

The asymptotic distribution of  $\hat{A_{n}}$  is not easily implemented.

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# **Goodness-of-Fit Methods for Complete and Right-Censored Data**

#### install.packages ("Gofcens") library(GofCens)

Graphical tools and goodness-of-fit tests for complete and right-censored data.

Goodness-of-fit test:

- KScens
- CvMcens
- ADcens
- chisqcens

Graphical tools:

- kmPlot
- probPlot
- cumhazPlot

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#### Example



•  $H_0: X \sim$  Weibull

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• Generates a plot that combines a **Kaplan-Meier survival curve** and a **parametric survival curve** in the same graph.



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# GofCens: probPlot

• Provides four types of probability plots: **P-P plot**, **Q-Q plot**, **Stabilised probability plot** and **Empirically Rescaled plot**.



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• Cumulative hazard plot to check if a certain distribution is an appropriate choice for the data.



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```
> KScens(colonsamp$time, colonsamp$status, distr = "weibull")
Distribution: weibull
KS Test results:
      A p-value F(ym) ym
  0.637 0.627 0.491 3238,000
Parameter estimates:
  shape scale
  0.762 4659.238
```
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- For the tests Cramér-von Mises, Anderson-Darling and Chi-square, the asymptotic distributions of goodness-of-fit statistics are difficult to obtain/implement in the presence of censored data.
- The p-values associated with these tests are obtained via **bootstrap** methods.

### <span id="page-20-0"></span>Bootstrap methods for right-censored data

 $H_0$ :  $F(t) = F_0(t; \theta)$ 

- **1** Observed data is utilized to estimate the parameter  $\theta$ , denoted as  $\hat{\theta}_p$ , using maximum likelihood estimation
- 2 Generation of B independent bootstrap samples of the same size  $(n)$  as the original data set:
	- Generation of survival times  $T_1^b, \ldots, T_n^b$  from the fitted distribution  $F_0(t; \hat{\theta}_n)$ .
	- Generation of censoring times  $C_1^b, \ldots, C_n^b$  from the nonparametric estimation of H obtained with the Kaplan-Meier estimator.
	- Generation of observed survival times  $Y_i^b = \min(T_i^b, C_i^b)$ , and event indicators  $\delta_i^b = \mathbf{1} \{ T_i^b \le C_i^b \}, i = 1, \dots n$

# <span id="page-21-0"></span>Bootstrap methods for right-censored data

- Maximum likelihood estimation of the parameter,  $\hat{\theta}_n^b$ , given  $(Y_i^b, \delta_i^b), i = 1, \ldots n.$
- Computation of the test statistic,  $(\hat{G}_n^{\hat{\theta}_n})_b$ .
- <sup>3</sup> Repetition of this process for many bootstrap samples (default B=999)
- $\bm{\Phi}$  Sequence of bootstrap statistics,  $(\hat{\bm{G}}_{{\bm{n}}}^{\hat{\theta}_{{\bm{n}}}})_{\bm{b}},~\bm{b}=1,\cdots,B$ , represents the empirical distribution of the statistic under the null hypothesis
- $\bm{s}$   $\bm{\rho}$  value is the proportion of bootstrap statistic values  $(\hat{\bm{G}}^{\hat{\theta}_n}_{{\bm{n}}})_{\bm{b}}$  greater than or equal to the observed statistic  $\hat{\mathsf{G}}_n$

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# What is the most correct test to apply? What does it depend on?

**Motivation** 

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## Simulation study



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#### Preliminary results

• Highly skewed Log-normal | Complete data



- Anderson-Darling and Cramér-von Mises have good power regardless of sample size.
- Kolmogorov-Smirnov has a good power for big sample size.
- Increase of sample size leads to increase of power.

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# Preliminary results

- Highly skewed Log-normal | 60 % censored data
- $H_0$  :  $X \sim$  Weibull



• Power results are not adequate regardless of the test applied.

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# Preliminary results

- Highly skewed Log-normal  $\vert$  60 % censored data
- $H_0: X \sim$  Logistic



- For large sample sizes, the tests produce good results.
- For a small sample size, Kolmogorov-Smirnov is the one that behaves better.
- <span id="page-29-0"></span>K. Langohr, M. Besalú, M. Francisco, G. Gómez, GofCens: Goodness-of-Fit Methods for Complete and Right-Censored Data, R package version 0.98 (2024).
- URL https://CRAN.R-project.org/package=GofCens T. R. Fleming, J. R. O'Fallon, P. C. O'Brien, D. P. Harrington, Modified kolmogorov-smirnov test procedures with application to arbitrarily
- right-censored data, Biometrics (1980) 607–625. J. A. Koziol, S. B. Green, A cramér-von mises statistic for randomly censored data, Biometrika 63 (3) (1976) 465–474.
- A. N. Pettitt, M. A. Stephens, Modified cramér-von mises statistics for censored data, Biometrika 63 (2) (1976) 291–298

# <span id="page-30-0"></span>Thank you for the attention! Gracias por la atención! Gràcies per l'atenció!