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Bonferroni and Holm approximations for Šidák and Holland–Copenhaver q-values

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Abstract. I describe the use of the Bonferroni and Holm formulas as approximations for Šidák and Holland–Copenhaver formulas when issues of precision are encountered, especially with q-values corresponding to very small p-values.

Keywords: st0300, parmest, qqvalue, smileplot, multproc, multiple-test procedure, familywise error rate, Bonferroni, Šidák, Holm, Holland, Copenhaver

1 Introduction

Frequentist q-values for a range of multiple-test procedures are implemented in Stata by using the package qqvalue (Newson 2010), downloadable from the Statistical Software Components (SSC) archive. The Šidák q-value for a p-value p is given by $q_{\rm sid} = 1 - (1-p)^m$, where m is the number of multiple comparisons (Šidák 1967). It is a less conservative alternative to the Bonferroni q-value, given by $q_{\rm bon} = \min(1, mp)$. However, the Šidák formula may be incorrectly evaluated by a computer to 0 when the input p-value is too small to give a result lower than 1 when subtracted from 1, which is the case for p-values of 10^{-17} or less, even in double precision. q-values of 0 are logically possible as a consequence of p-values of 0, but in this case, they may be overliberal. This liberalism may possibly be a problem in the future, given the current technology-driven trend of exponentially increasing multiple comparisons and the human-driven problem of ingenious data dredging. I present a remedy for this problem and discuss its use in computing q-values and discovery sets.

2 Methods for q-values

The remedy used by the SSC packages qqvalue and parmest (Newson 2003) is to substitute the Bonferroni formula for the Šidák formula for such small *p*-values. This works because the Bonferroni and Šidák *q*-values converge in ratio as *p* tends to 0. To prove this, I show that for $0 \le p < 1$,

 $dq_{\rm bon}/dp = m$ and $dq_{\rm sid}/dp = m(1-p)^{m-1}$

and that the Šidák/Bonferroni ratio of these derivatives is $(1-p)^{m-1}$, which is 1 if p = 0. By L'Hôpital's rule, it follows that the ratio $q_{\rm sid}/q_{\rm bon}$ also tends to 1 as p tends to 0.

A similar argument shows that the same problem exists with the q-values output by the Holland–Copenhaver procedure (Holland and Copenhaver 1987). If the m input p-values, sorted in ascending order, are denoted p_i for i from 1 to m, then the Holland– Copenhaver procedure is defined by the formula

$$s_i = 1 - (1 - p_i)^{m-i+1}$$

where s_i is the *i*th *s*-value. (In the terminology of Newson [2010], *s*-values are truncated at 1 to give *r*-values, which are in turn input into a step-down procedure to give the eventual *q*-values.) The remedy used by **qqvalue** here is to substitute the *s*-value formula for the procedure of Holm (1979), which is

$$s_i = (m - i + 1)p_i$$

whenever $1 - p_i$ is evaluated as 1. This also works because the two s-value formulas converge in ratio as p_i tends to 0. Note that the Holm procedure is derived from the Bonferroni procedure by using the same step-down method as is used to derive the Holland–Copenhaver procedure from the Šidák procedure.

3 Methods for discovery sets

The SSC package smileplot (Newson and the ALSPAC Study Team 2003) also implements a range of multiple-test procedures by using two commands, multproc and smileplot. However, instead of outputting q-values, smileplot outputs a corrected critical p-value threshold and a corresponding discovery set, defined as the subset of input p-values at or below the corrected critical p-value. The Šidák-corrected critical p-value corresponding to an uncorrected critical p-value p_{unc} is given by $c_{sid} = 1 - (1 - p_{unc})^{1/m}$ and may be overconservative if wrongly evaluated to 0. In this case, the quantity that might be wrongly computed as 1 is $(1 - p_{unc})^{1/m}$. When this happens, smileplot substitutes the Bonferroni-corrected critical p-value $c_{bon} = p_{unc}/m$. However, this is a slightly less elegant remedy in this case because the quantity $(1 - p_{unc})^{1/m}$ is usually evaluated to 1 because m is large and not because p_{unc} is small.

To study the behavior of the Bonferroni approximation for large m, we define $\lambda = 1/m$ and note that

$$dc_{\rm bon}/d\lambda = p_{\rm unc}$$
 and $dc_{\rm sid}/d\lambda = -\ln(1-p_{\rm unc})(1-p_{\rm unc})^{\lambda}$

implying (by L'Hôpital's rule) that in the limit, as λ tends to 0, the Šidák/Bonferroni ratio of the two derivatives (and therefore of the two corrected thresholds) tends to $-\ln(1-p_{\rm unc})/p_{\rm unc}$. This quantity is not as low as 1 but is 1.150728, 1.053605, 1.025866, and 1.005034 if $p_{\rm unc}$ is 0.25, 0.10, 0.05, and 0.01, respectively. Therefore, the Bonferroni approximation in this case is still slightly conservative for a very large number of multiple comparisons over a range of commonly used uncorrected critical *p*-values, but is less conservative than the value of 0, which would otherwise be computed.

This argument is easily generalized to the Holland–Copenhaver procedure. In this case, smileplot initially calculates a vector of m candidate critical p-value thresholds by using the formula

$$c_i = 1 - (1 - p_{\text{unc}})^{1/(m-i+1)}$$

for *i* from 1 to *m* and selects the corrected critical *p*-value corresponding to a given uncorrected critical *p*-value from these candidates by using a step-down procedure. If the quantity $(1 - p_{\text{unc}})^{1/(m-i+1)}$ is evaluated as 1, then smileplot substitutes the corresponding Holm critical *p*-value threshold

$$c_i = p_{\rm unc}/(m-i+1)$$

which again is conservative as m - i + 1 becomes large (corresponding to the smallest *p*-values from a large number of multiple comparisons), but is less conservative than the value of 0, which would otherwise be computed.

Newson (2010) argues that q-values are an improvement on discovery sets because, given the q-values, different members of the audience can apply different input critical p-values and derive their own discovery sets. The technical issue of precision presented here may be one more minor reason for preferring q-values to discovery sets.

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5 References

- Holland, B. S., and M. D. Copenhaver. 1987. An improved sequentially rejective Bonferroni test procedure. *Biometrics* 43: 417–423.
- Holm, S. 1979. A simple sequentially rejective multiple test procedure. Scandinavian Journal of Statistics 6: 65–70.
- Newson, R. 2003. Confidence intervals and p-values for delivery to the end user. Stata Journal 3: 245–269.
- Newson, R., and the ALSPAC Study Team. 2003. Multiple-test procedures and smile plots. *Stata Journal* 3: 109–132.
- Newson, R. B. 2010. Frequentist q-values for multiple-test procedures. Stata Journal 10: 568–584.
- Šidák, Z. 1967. Rectangular confidence regions for the means of multivariate normal distributions. *Journal of the American Statistical Association* 62: 626–633.

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Roger B. Newson is a lecturer in medical statistics at Imperial College London, UK, working principally in asthma research. He wrote the parmest, qqvalue, and smileplot Stata packages.