# Stata Technical Bulletin

A publication to promote communication among Stata users

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stata52	Origin/noOrigin option added to sts graph command	
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The sts graph command now plots the Kaplan-Meier survival curve starting at t = 0; see [R] st sts graph. This is now the default. Specifying the new option noorigin returns to the previous behavior of having the graph start at the first observed exit time (failure or censoring). All other options for this command are unchanged.

# Syntax

```
sts graph [if exp] [in range] [, by(varlist) strata(varlist) adjustfor(varlist) nolabel failure
    gwood level(#) lost enter separate tmin(#) tmax(#) xasis yasis noborder noshow ]
    noorigin graph_options ]
```

The noorigin option is new. See [R] st sts graph for a description of the other options.

noorigin sets the earliest time plotted to correspond to the first failure or censoring time. If this option is not specified, zero is the earliest time plotted.

# Output

Using the cancer data distributed with Stata, we will plot the Kaplan-Meier product-limit estimate for patients on drug 2, and we will repeat the plot specifying the noorigin option which will cause the plot to begin at studytim = 6, the first exit time for patients on drug 2.

. use cancer.dta (Patient Survival in Drug Trial)						
. stset stud	lytim died					
(output omitted). sort study	,					
. list if dr	ug==2, no	obs				
<pre>studytim died drug age 6 1 2 67 6 0 2 65 7 1 2 58 9 0 2 56 10 0 2 49 (output omitted) . sts graph if drug==2 ( See Figure 1 below)</pre>						
. sts graph (See Figure 2)	0	2, noorigin				

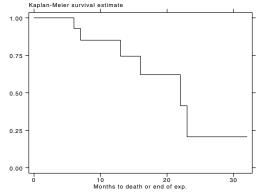


Figure 1. KM curve for patients on drug 2

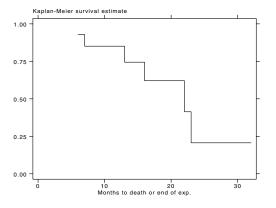


Figure 2. KM curve for patients on drug 2; noorigin specified

dm54	Capturing comments from data dictionaries
	Captaining commonte nom data dictionance

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Data dictionaries are an effective way to import and export data: The data are transported in an ASCII text file (a reliable mechanism on any reasonable computing platform), and the dictionary itself provides a clear description of the contents of the data file. (See [R] **infile (fixed format)** for details.) More importantly, dictionaries can be freely commented: any line with the character '\*' in the first column will be ignored by the **infile** command when it uses the dictionary to read the data. Comments can thus be used to document the source of the data, or to explain details regarding the collection of the data—information that ought to remain attached to the data. This insert presents a command (infilen) that creates a Stata dataset by infiling from a fixed format data file, captures commentary from the data dictionary, and stores those comments as notes; see [R] **notes**.

To illustrate, consider the file mammals.dat (included with this insert) which begins thus:

```
dictionarv {
*! Data originally published by Allison, T. & Cicchetti, D. V. (1976)
*! <Science, 194, 732-734>, as part of a study of correlates of sleep in
*! mammals; a subset (the first 3 variables below) appears in Table 6.6 of
*! Weisberg (1985) < Applied Regression, 2nd ed.; NY: Wiley; pp. 144-145>.
* 1
*! The data show average values for 62 species of mammals; note that some
*! values are missing. The last 3 variables below (is prey, expos_sl, and
*! indanger) are ordered categories, where 1 = least and 5 = most with
*! respect to the attribute in question. Note also that slow wave sleep
*! (swav_sl) is known as `non-dreaming' sleep, whereas paradoxical sleep
  (pdox_sl) is known as `dreaming' sleep.
*!
   str25
         species
                     "Mammal species"
          body_wt
                     "Body Weight (kg)"
                     "Brain Weight (g)"
          brain wt
          swav_sl
                     "Slow wave sleep (hrs/day)"
          pdox_sl
                     "Paradoxical sleep (hrs/day)"
                     "Total sleep (hrs/day)"
          totsleep
                     "Maximum life span (yrs)"
          lifespan
                     "Gestation time (days)"
          gestate
   byte
          is_prey
                     "Likely to be preyed upon?"
   byte
          expos_sl
                     "Exposed during sleep?"
          indanger
                     "In danger from other animals?"
   byte
}
                                                       3.3 38.6 645 3 5 3
         "African elephant"
                              6654
                                     5712
"African giant pouched rat"
                                 1
                                      6.6
                                            6.3
                                                   2
                                                       8.3 4.5 42 3 1 3
 (output omitted)
```

The command

. infile using mammals.dat

will create a Stata dataset with 11 usefully labeled variables, but the information contained in the lines beginning with the characters '\*!' will be left behind. By contrast, the command

. infilen using mammals.dat

creates the same Stata dataset but also imbeds the lines marked with '\*!' as 11 notes.

The syntax of infilen is

infilen using dilename [if exp] [in range] [, automatic using(filename<sub>2</sub>) clear <u>n</u>lines(#)]

The option nlines controls the number of data dictionary lines that will be scanned for comments; the default value of # is 50. Otherwise, the syntax of infilen is exactly that of infile for reading data with a dictionary; see [R] infile (fixed format) for details.

In fact, the command

. infilen using mammals.dat

first issues the command

. infile using mammals.dat

Then, infilen re-reads the data dictionary in mammals.dat, searching for lines where the string '\*!' appears in the first two columns. This marks a special type of comment, the kind that if found in an ado-file, will be echoed to the screen by which; see [R] which. infilen captures such comments and then does the equivalent of

. note: text of comment

for each '\*!' comment. Thus, continuing our example, after the infilen command there will be 11 notes attached to the current dataset:

```
. notes
_dta:
 1. Data originally published by Allison, T. & Cicchetti, D. V. (1976)
 2. <Science, 194, 732-734>, as part of a study of correlates of sleep in
 3. mammals; a subset (the first 3 variables below) appears in Table 6.6 of
  4.
     Weisberg (1985) < Applied Regression, 2nd ed.; NY: Wiley; pp. 144-145>.
 5.
 6. The data show average values for 62 species of mammals; note that some
  7.
     values are missing. The last 3 variables below (is_prey, expos_sl, and
 8. indanger) are ordered categories, where 1 = least and 5 = most with
 9. respect to the attribute in question. Note also that slow wave sleep
10.
     (swav_sl) is known as 'non-dreaming' sleep, whereas paradoxical sleep
      (pdox_sl) is known as 'dreaming' sleep.
11.
```

# Remarks

Ordinary comments (those that begin with '\*' only) are simply ignored, while every comment with '\*!' in the first two columns is stored in a note. Blank comments are often used to make commentary more readable; for compatibility with the notes command, such comments are rendered non-blank by saving the single character '.'. Note 5 (above) provides an example.

Instances of the left-quote (``) character will be translated into right-quotes (`'); otherwise, some part of the comment will be interpreted as a macro expansion. Unfortunately, there is no way to perform a similar translation of the double-quote character (`"): Comments to be saved as notes may not contain `"', and its presence will produce an unhelpful error message. (On the other hand, the characters ``' and `"' have no special status in ordinary comments.)

A maximum of 80 characters following the string '\*!' will be captured from comments.

# Acknowledgment

This project was supported by a grant R01-MH54929 from the National Institute on Mental Health to Michael P. Carey.

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The problem of testing the equality of several variances arises in many areas. There are a wide variety of tests available for testing equality of variances of normal populations. In Stata, only Bartlett's (1937) test for equality of variances is available in the oneway command.

The grvar command performs a graphical display for testing the equality of variances of normal populations. Rao and Krishna (1997) following Ott's (1967) graphical procedure for testing the equality of means (ANOM) have developed this method. This graph tests equality of two or more variances from normal populations, simultaneously demonstrating statistical (the real presence of an effect) and engineering significance (the magnitude of seriousness of the effect). This graphical method could be more useful for non-statisticians than classical tests for equal variances, becoming a good supplement to the oneway command.

## Graphical method procedure

Consider k independent samples, the *i*th of which is of size  $n_i$ , obtained from k normal populations with means  $\mu_i$  and variances  $\theta_i^2$ . We would like to test the null hypothesis  $\theta_i^2 = \theta^2$ , for all *i*, against the alternative that at least one equality does not hold. Then, we should follow four steps:

1. Calculate the variance of the *i*th sample  $(S_i^2)$ , and then  $S_i^{2/3}$ , for i = 1, ..., k.

2. Calculate: 
$$A_i = \frac{(9n_i - 11)}{9(n_i - 1)}, \qquad C_i = \frac{18(n_i - 1)}{(9n_i - 11)^2}, \qquad N = \sum_{i=1}^k n_i, \qquad t_i = \frac{S_i^{2/3}}{A_i},$$
  
 $\bar{t}_w = \frac{1}{N} \sum_{i=1}^k n_i t_i, \qquad SE(t_i - \bar{t}_w) = \frac{\bar{t}_w}{N} \sqrt{N(N - 2n_i)C_i + \sum_{i=1}^k n_i^2 C_i}$ 

- 3. Calculate the lower and upper decision lines (LDL and UDL) for the comparison of each of the  $t_i$  terms: LDL =  $\bar{t}_w z_{a/2k}SE(t_i \bar{t}_w)$ , UDL =  $\bar{t}_w + z_{a/2k}SE(t_i \bar{t}_w)$ , where  $z_{a/2k}$  is the upper (a/2k) percentage point of the standard normal distribution.
- 4. Finally, plot  $t_i$  against LDL and UDL.

Derivations of the formulas are given in the appendix of Rao and Krishna (1997).

If any of the points plotted are outside the decision lines, we can reject the null hypothesis and conclude that there are statistically significant differences among the population variances.

# Syntax

The grvar command works on a dataset containing the estimated standard deviation, sd, and the sample size, n, for each sample or study. The syntax is as follows:

grvar *sd n* [if *exp*] [in *range*] [, id(*strvar*) level(#) *graph\_options*]

# Options

id(strvar) is used to label the studies. If the data contain a labeled numeric variable, it can also be used.

level (#) specifies the level of significance of the test, in percent. The default is level(5) (sic).

graph\_options are any of the options allowed with graph, twoway except xlabel(), t2(), b2(), yline(), symbol(), pen(), and connect().

#### Example

The grvar command has been tested using the example supplied by Rao and Krishna (1997). We have eight samples we call A, B, C, D, E, F, G, and H of lengths of the third molar of eight species of the Condylarth Hyposodus (Sokal and Rohlf 1969, 370–371).

```
. describe
Contains data from sokal.dta
  obs:
                     8
vars:
                     4
                    88
size:
                            %8.0g
                    byte
                                                     sample number
   1. sample
   2. sampleid
                    str1
                            %9s
                                                     sample identification
   3. n
                            %8.0g
                                                     sample size
                    byte
   4. sd
                    float
                           %9.0g
                                                     standard deviation
. list, noobs
        sample
                  sampleid
                                                sd
                                    n
                                         .2658947
                                   18
                         A
             1
                                         .3803945
             2
                         В
                                   13
             3
                         С
                                   17
                                          .153948
             4
                         D
                                   16
                                         .2891366
             5
                         Е
                                    8
                                          .4678675
             6
                         F
                                   11
                                         .4207137
             7
                         G
                                   10
                                         .2812472
             8
                                   10
                                         .4828043
                         Н
 grvar sd n, id(sampleid)
                     .698497
                                                 D
                                                                      6
                     244335
                                                        5
                                                               6
                                                Sample number
```

Figure 1. Graphical display to test equality of variances

We observe from Figure 1, that sample C is allocated outside its lower decision line. Hence we conclude that the population variances are not equal. Sokal and Rohlf (1969) using Bartlett's test at the 5% level obtained the same conclusion. Also Rao and Krishna (1997) got the same result using the Neyman–Pearson test, the Lehmann test, the Q test, the Bartlett 3 test, the Lehmann 2 test, and the Samiuddin–Atiqullah test with their respective critical values at the 5% level.

## Individual or frequency records

As with other Stata commands, grvar works on data contained in frequency records, one for each sample or study. If we have primary data, that is individual records, we must convert to frequency records using the collapse and byvar Stata commands.

## References

Bartlett, M. S. 1937. Properties of sufficiency and statistical tests. Proceedings of the Royal Statistical Society, Series A 160: 268-282.

Ott, E. R. 1967. Analysis of means-a graphical procedure. Industrial and Quality Control 24: 101-109.

Rao, C. V. and S. H. Krishna. 1997. A graphical method for testing the equality of several variances. *Journal of Applied Statistics* 24: 279–287. Sokal, R. R. and J. F. Rohlf. 1969. *Biometry*. San Francisco, CA: W. H. Freeman.

sbe16.1	New syntax and output for the meta-analysis command
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Stephen Sharp, London School of Hygiene and Tropical Medicine, stephen.sharp@lshtm.ac.uk Jonathan Sterne, United Medical and Dental Schools, UK, j.sterne@umds.ac.uk

The command meta, which performs the statistical methods involved in a systematic review of a set of individual studies, reporting the results in text and also optionally in a graph, was released in STB-38 (Sharp and Sterne 1997). This re-release contains a more flexible syntax than the earlier version and new-style output.

## Syntax

The command meta works on a dataset containing an estimate of the effect and its variability for each study. The syntax is

meta {theta | exp(theta)} {se\_theta | var\_theta | ll ul [cl]} [if exp] [in range]
 [, var ci eform print ebayes level(#) graph(string) id(varname)
 fmult(#) boxysca(#) boxshad(#) cline ltrunc(#) rtrunc(#) graph options]

With this new syntax, the user provides the effect estimate as *theta* (i.e., a log risk ratio, a log odds ratio, or other measure of effect). Likewise, the user supplies a measure of *theta*'s variability (i.e., its standard error, *se\_theta*, or its variance, *var\_theta*). Alternatively, the user provides *exp(theta)* (e.g. a risk ratio or odds ratio) and its confidence interval, (*ll*, *ul*). The options remain the same as in the original version of meta. Type help meta for details.

# **Required input variables**

theta se_theta	the effect estimate the corresponding standard error
or	
<i>theta</i> <i>var_theta</i> NB: (var option must be included)	the effect estimate the corresponding variance
or	
exp(theta) ll ul cl NB: (ci option must be included)	the risk (or odds) ratio the lower limit of the risk ratio's confidence interval the upper limit of the risk ratio's confidence interval optional (see below)

#### Optional input variable

cl contains the confidence level of the confidence interval defined by ll and ul. If cl is not provided, the procedure assumes that each confidence interval is at the 95% level. cl allows the user to provide the confidence level, by study, when the confidence interval is not at the default level. cl can be specified with or without a decimal point. For example, .90 and 90 are equivalent and may be mixed (i.e., 90, .95, 80, .90 etc.).

#### Example of new syntax and output

Below we illustrate the new-style command syntax and output from meta applied to the example in Sharp and Sterne (1997), based on nine randomized clinical trials of the use of diuretics for pre-eclampsia in pregnancy (Collins et al. 1985).

```
use diuretic. clear
(Diuretics and pre-eclampsia)
. describe
Contains data from diuretic.dta
 obs:
                  9
                                               Diuretics and pre-eclampsia
vars:
                  6
size:
                189
                        %9.0g
  1. trial
               byte
                                    trlab
                                            trial identity number
  2. trialid
                 str8
                        %9s
                                               trial first author
  3. nt
                        %9.0g
                                               total treated patients
                 int
  4. nc
                 int
                        %9.0g
                                               total control patients
                        %9.0g
  5. rt
                  int
                                               pre-eclampsia treated
                        %9.0g
  6. rc
                 int
                                               pre-eclampsia control
```

With the earlier version of meta it was necessary to calculate the estimated effect and standard error on the *log* scale. With the re-release, the user can specify either the odds ratio (on either the original or log scale), the standard error, the variance, or the confidence limits. So for example, suppose the odds ratio and 90% confidence limits are calculated.

```
. gen or=(rt/(nt-rt))/(rc/(nc-rc))
. gen 190=or/exp(1.645*sqrt((1/rc)+(1/(nc-rc))+(1/rt)+(1/(nt-rt))))
. gen u90=or*exp(1.645*sqrt((1/rc)+(1/(nc-rc))+(1/rt)+(1/(nt-rt))))
. 1 190 or u90, noobs
     190
                 or
                           u90
 .5404617
           1.042735
                      2.011792
           .3970588
 .2257975
                      .698217
  .162642
           .3255814
                       .651758
  .093229
           .2291667
                      .5633156
 .1426823
           .2488202
                      .4339115
 .6090319
           .7431262
                      .9067448
 .4347324
           .7698574
                      1.363322
 .7602696
           2.970588
                      11.60693
.7458201
           1.144886
                     1.757481
. gen cl=90
. meta or 190 u90 cl, eform ci
Meta-analysis (exponential form)
         Pooled
                  95% CI
                                    Asymptotic
                                                    No. of
Method
            Est
                  Lower
                          Upper z_value p_value
                                                    studies
Fixed 0.672
                  0.564
                          0.800
                                  -4.454
                                            0.000
                                                       9
                  0.400
                          0.889
                                  -2.537
Random
          0.596
                                            0.011
Test for heterogeneity: Q= 27.260 on 8 degrees of freedom (p= 0.001)
Moment-based estimate of between studies variance = 0.230
```

Note that although in this example the odds ratios and confidence intervals had to be calculated before the meta command could be used, in practice such variables may already have been created in the dataset for other purposes, and the new syntax saves the user from having to generate the log odds ratio and standard error. If the effect estimate is a difference in means, the same alternatives for specifying the variability are available.

## Graphics

The graphics options remain as in the earlier version of the command (Sharp and Sterne 1997).

# Acknowledgment

We are grateful to Thomas Steichen for his helpful suggestions regarding the command syntax and output.

## References

Collins, R., S. Yusuf, and R. Peto. 1985. Overview of randomised trials of diuretics in pregnancy. *British Medical Journal* 290: 17–23. Sharp, S. and J. Sterne. 1997. sbe16: Meta-analysis. *Stata Technical Bulletin* 38: 9–14.

sbe21	Adjusted population attributable fractions from logistic regression
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The population attributable fraction (or risk) is an epidemiological measure which attempts to quantify the proportion of disease incidence which is due to a particular factor (or "exposure"). Implicit in this measure are very strong assumptions about causality, namely that removing the exposure from the population would reduce disease incidence. However, where this assumption is met the attributable fraction can provide useful insights into the important causes of disease and thereby inform intervention strategies.

The usual measure of attributable fraction (such as that reported by Stata's epitab cc command) is an unadjusted measure which can be misleading in the presence of confounding. Estimating the attributable fraction (AF) from within a logistic regression framework enables confounders to be taken into account and allows estimation of the summary attributable fraction for a set of exposures. This might be of interest to those planning an intervention "package".

aflogit recovers estimates of the population attributable fraction for those terms specified by the user from the most recent unconditional logistic regression model. It also reports a summary attributable fraction for all the terms combined. Asymptotic standard errors and 95% confidence intervals are calculated and reported. Note that aflogit may also be used following Poisson regression (where the attributable fraction is interpreted as the proportion of the disease rate which is due to the exposure), but cannot be used after conditional logistic regression (clogit).

Negative estimates of the attributable fraction correspond to protective effects (relative risk < 1) where the interpretation is the proportion increase expected in the number with the disease if the protective exposure were absent from the population. Perhaps confusingly this is not the usual definition of the preventable fraction or the prevented fraction (see Last 1983).

#### Background

The unadjusted population attributable fraction is given by

$$AF = \frac{\Pr(\text{disease}) - \Pr(\text{disease}|\text{not exposed})}{\Pr(\text{disease})}$$

which is equivalent to

Observed no. ill

This can be rearranged to give

$$AF = Pr(exposed|disease) \left(1 - \frac{1}{RR}\right)$$

where RR is the relative risk. So all that is needed to estimate the AF is the distribution of exposure among the ill, Pr(exposed | disease), and the relative risk. The distribution of exposure among the ill is assumed to apply to the population from which the sample was taken. Therefore, any sampling procedure which does not randomly sample those with the disease will invalidate estimation of the AF. Under the rare disease assumption, the relative risk is approximated by the odds ratio and so case-control data can be used to estimate the AF. This is the formula the epitab cc command uses.

Notice that the attributable fraction increases both as the exposure becomes more common and as the relative risk becomes larger. In addition, uncertainty in the estimated AF will come not only from the uncertainty in the relative risk but also from the distribution of exposure. This means that a risk factor which is significantly different from one may not necessarily have an attributable fraction which is significantly bigger than zero.

The principle behind using logistic regression to estimate attributable fractions is to fit the required model and then calculate the number of cases predicted by the model. This is done by generating the predicted probability of disease for each individual in the model (using the predict command) and then summing over all individuals. If the model has been fitted to the entire dataset then this should equal the observed number of cases of disease in the study. Next the exposure effect is "removed" from the dataset by resetting exposure covariates to zero (or some reference level for continuous variables). The predict command is used again to generate the predicted probability of disease for each individual with the new covariate values but under the same logistic model. Summing these probabilities gives the number of cases of disease one would expect if the exposure were absent from the population. Calculating the AF is then a simple matter of applying the second equation above.

More formally, the adjusted attributable fraction is given by

$$\widehat{\lambda}_A = 1 - \frac{n'r_z}{n'r_x}$$

where n is the column vector of sample sizes,  $n_i (\equiv 1$  where records are on individuals), and  $r_x$  and  $r_z$  are vectors of predicted probabilities from the model under the original covariates (x) and the modified covariates (z) respectively.

An asymptotic variance formula is given in Greenland and Drescher (1993), which they recommend be applied on the log(1 - AF) scale. This is most helpfully considered as

$$\operatorname{var}(\widehat{\lambda}) = \left(\frac{E}{O}\right)^2 \left(\frac{\operatorname{var}_E}{E^2} + \frac{\operatorname{var}_O}{O^2} - \frac{2\operatorname{cov}(O, E)}{OE}\right)$$

where O and E refer to the number of observed and expected cases of disease respectively.

A different approach must be taken for case–control data since the Pr(disease) is fixed by the study design (0.5 if one control per case is used). Bruzzi et al. (1985) explain this approach, although it is also presented by Greenland and Drescher (1993) with some minor modifications. Essentially the method is to generalize the third equation to J strata (or levels of exposure) which gives

$$AF = 1 - \sum_{j=0}^{J} \frac{\rho_j}{RR_j}$$

where  $\rho_j = \Pr(\text{exposure group } j \mid \text{disease})$  and  $RR_j$  is the relative risk comparing stratum j with stratum 0, the baseline group. The  $RR_j$  may of course be approximated by  $OR_j$  if the rare disease assumption is justified. This simplifies to the third equation when there are only two strata (exposed and not exposed)

$$\begin{aligned} AF &= 1 - \left(\frac{\Pr(\text{not exposed}|\text{disease})}{1} + \frac{\Pr(\text{exposed}|\text{disease})}{\text{RR}}\right) \\ &= 1 - 1 + \Pr(\text{exposed}|\text{disease}) - \frac{\Pr(\text{exposed}|\text{disease})}{\text{RR}} \\ &= \Pr(\text{exposed}|\text{disease}) \left(1 - \frac{1}{\text{RR}}\right) \end{aligned}$$

The above generalized equation gives the summary attributable fraction for all the factors combined, that is, the proportion of the disease which is due to all the exposures under consideration. In order to estimate the effect of one risk factor while adjusting for other covariates, we modify the  $RR_j$  to reflect the additional risk which the risk factor of interest contributes to stratum j. Strata in which the risk factor does not appear therefore have RR = 1. The  $RR_j$  for strata including the risk factor is estimated by the adjusted odds ratio from the logistic regression model (making the rare disease assumption). Providing the risk factor of interest is dichotomous and is not involved in any interactions, the estimate of the adjusted AF reduces to substituting the adjusted odds ratio into the third equation. In the notation used by Greenland and Drescher (1993), the generalized equation is

$$\lambda_A = 1 - \hat{
ho}' \hat{s}$$

where  $\hat{\rho}$  is a vector containing the  $\hat{\rho}_j$  and  $\hat{s}$  is a vector of  $\frac{1}{\widehat{RR}_j}$ .

The asymptotic variance of this measure has no simple form and aflogit uses just the first summand in Greenland and Drescher's formula,  $var(\hat{\lambda}) \approx D'_{\theta}CD_{\theta}$ , which they report gives results nearly identical to those based on the entire formula.

#### Syntax

The command aflogit may only be used after a logistic or poisson regression model has been fitted. The syntax is

fweights and aweights are allowed.

Without arguments, aflogit reports attributable fractions for all terms in the model. aflogit tries to pick up the same weights, if and in clauses as used by the regression model, but users should manually specify these after the logit, blogit, and poisson commands.

# Options

- cc indicates that the data come from an unmatched case-control study and the estimate will be based on the adjusted odds ratios. The default is to assume the data come from a cross-sectional or cohort study and base the estimate of the attributable fraction on the predicted probabilities. It is imperative to select this option if you have case-control data since the results might be seriously misleading if you use the predicted probabilities.
- reference determines the reference level (the non-exposure category) for each term in the model. By default this is zero for every term. For continuous variables it may be desirable to set the reference level to a nonzero value (e.g. the mean for a variable such as blood pressure).
- level (#) specifies the significance level for confidence intervals of the coefficients.

#### Example: unmatched case-control data

Jacques Benichou (1991) calculated population attributable fraction estimates for a case–control study of oesophageal cancer carried out by Tuyns et al. (1977). The main interest was in the proportion of cases attributable to excess alcohol consumption. Various logistic regression models were fitted to allow for confounders (smoking and age) and for interactions between alcohol consumption and the confounders. Alcohol consumption (the "exposure") was collected as a four-level factor but was initially used as a dichotomous factor (0 to 79 grams per day and 80 or more grams per day).

```
. use tuvns, clear
(Desophageal cancer and alcohol)
. describe
Contains data from tuyns.dta
                  59
  obs:
                                                 Desophageal cancer and alcohol
                                                 17 Sep 1997 17:07
 vars:
                   7
                 649 (100.0% of memory free)
 size:
   1. alcohol
                  byte
                         %9.0g
                                      alcl
                                                 g/day
                         %9.0g
   agegp
                  byte
                                      agel
                                                 Age group
                         %9.0g
                                                 g/day
   3. smoke
                  byte
                                      smokel
                         %9.0g
   4. case
                  byte
                                                 Cancer case?
   5. n
                  byte
                         %8.0g
                                                 Frequency
   6. alc80
                                                 Alcohol 0-79 vs 80+
                  byte
                         %9.0g
                         %9.0g
                                                 Alcohol 0-39 vs 40+
   7. alc40
                  byte
Sorted by: case
. cc case alc80 [fw=n]
                   Alcohol 0-79 vs 80+
                                                         Proportion
                     Exposed Unexposed
                                                 Total
                                                           Exposed
           Cases
                          96
                                      104
                                                   200
                                                            0.4800
                                                   775
                         109
                                      666
        Controls
                                                            0.1406
                                           ____
                                                   ----
           Total
                         205
                                      770
                                                   975
                                                            0.2103
                           Pt. Est.
                                              [95% Conf. Interval]
      Odds ratio
                           5.640085
                                              4.003217
                                                           7 94673
                                                                     (Cornfield)
 Attr. frac. ex.
                            .8226977
                                              .7502009
                                                          .8741621
                                                                    (Cornfield)
                            .3948949
 Attr. frac. pop
                                           chi2(1) =
                                        110.26 Pr>chi2 = 0.0000
```

Thus using the cc command we estimate that 39% of the oesophageal cancers were attributable to alcohol consumptions above 80 g/day. Note there is no confidence interval around this estimate. We can obtain the same result using logistic regression and aflogit but with the added benefit of a confidence interval.

. logis	tic case	alc80 [f	w=n]				
Logit E	stimates					Number of ob chi2(1) Prob > chi2	= 96.43
Log Lik	elihood =	-446.52	782			Pseudo R2	
cas	e   Odds	Ratio	Std. Err.	z	P> z	[95% Conf.	Interval]
alc	80  5.6	40085	.9883491	9.872	0.000	4.00059	7.951467
. aflog							
-	ion attri ntrol dat		fraction from )	n logisti	c regres	sion	
Using w	eights [f	weight=	n]				
Term	Ref.	A.F.	s.e.	[95%	Conf. I	nt.]*	
			0.0409				
			0.0409				
* CI ca	lculated	on log(1	-AF) scale				

The logistic regression framework also allows us to extend the model to adjust for potential confounders (age-group and smoking in this example) and even to allow for interactions between the exposure and confounders; for instance, excess alcohol consumption may have a worse effect depending on age-group. First, allowing for potential confounders we have

i.agegp i.smoke			(naturall) (naturall)	y coded; y coded;	lagegp_0 omitte Ismoke_0 omitte	
Logit Est	imates				Number of ob	os = 975
-					chi2(9)	= 232.84
					Prob > chi2	
Log Likel	ihood = -378	3.32187			Pseudo R2	= 0.2353
			z	P> z	[95% Conf.	. Interval]
	4.8823		8.213	0.000	3.3442	7.128055
Iagegp_1	11.8125	9.13967	3.191	0.001	2.592679	53.8195
Iagegp_2	27.8718	20.35787		0.000	6.659488	116.6518
Ismoke_1	6.390786		2.297			31.11572
	2.33e-0		-34.902			
					.0460605	
					7889856	
		.1856753	-1.797	0.072	.0426983	1.146182
IaXs_2_2	2.20e+0	· .	•	•	•	•
Note: 33	failures and	l O successes	completely	determi	ned.	
. aflogit	alc80, cc		- •			
-	n attributal rol data (n=	ole fraction : =975)	from logist:	ic regre	ssion	
Using wei	ghts [fweigh	nt= n]				
Term	Ref. A	.F. s.	e. [95;	Conf.	Int.]*	
alc80	0 0.3	3817 0.04	442 0.2	2887 0	.4625	
TOTAL	0.3	3817 0.04	442 0.2	2887 0	.4625	
* CI calc	ulated on lo	g(1-AF) scale	e			

\* CI calculated on log(1-AF) scale

The odds ratio on alcohol changed little (from 5.6 to 4.9) when allowing for smoking and age and so the adjusted attributable fraction is very similar (38%). Now allowing alcohol consumption to interact with age-group, we have

```
. xi: logistic case i.agegp*alc80 i.agegp*i.smoke [fw=n]
i.agegp Iagegp_0-2 (naturally coded; Iagegp_0 omitted)
i.agegp*alc80 IaXal1_# (coded as above)
i.smoke Ismoke_0-2 (naturally coded; Ismoke_0 omitted)
i.agegp*i.smoke IaXs_#-# (coded as above)
Note: Iagegp_1 dropped due to collinearity.
Note: Iagegp_2 dropped due to collinearity.
```

Logit Estimates					Number of ob chi2(11) Prob > chi2	s = 975 = 233.13 = 0.0000
Log Likel:	ihood = -378	Pseudo R2	= 0.2356			
case	Odds Ratio	Std. Err.	z	P> z	[95% Conf.	Interval]
Iagegp_1	12.75294	10.61766	3.058	0.002	2.494186	65.20659
Iagegp_2	31.46491	24.56128	4.418	0.000	6.813728	145.3008
alc80	6.527618	4.431543	2.763	0.006	1.725367	24.69608
IaXalc_1	.8002673	.6210458	-0.287	0.774	.1748485	3.662759
IaXalc_2	.7024653	.5052614	-0.491	0.623	.1715466	2.876521
Ismoke_1	6.226884	5.062682	2.249	0.024	1.265357	30.6428
Ismoke_2	2.14e-07	1.25e-07	-26.312	0.000	6.83e-08	6.72e-07
IaXs_1_1	.275004	.2492789	-1.424	0.154	.0465332	1.625232
IaXs_1_2	3.58e+07				•	•
IaXs_2_1	.22786	.1924345	-1.751	0.080	.0435315	1.192705
IaXs_2_2	2.38e+07	1.74e+07	23.318	0.000	5716679	9.94e+07

Note: 33 failures and 0 successes completely determined.

. aflogit alc80 IaXalc\*, cc

Population attributable fraction from logistic regression Case-control data (n=975) Using weights [fweight= n] Term Ref. A.F. s.e. [95% Conf. Int.]\*

alc80	0	0.4065	0.0630	0.2693	0.5179
IaXalc_1	0	-0.0312			
IaXalc_2	0	-0.1398	•		
TOTAL		0.3803	0.0445	0.2866	0.4616
			-		

\* CI calculated on log(1-AF) scale

There appears to be little evidence of an interaction between age and alcohol consumption and so the attributable fraction is unchanged. The negative values of the attributable fraction indicate that removing an exposure would make the expected number of cases larger than that observed. Hence, removing the slight protection afforded by being in the oldest age-group would result in a 14% increase in the observed number of cases of oesophageal cancer in the population. Note that algebraically the population attributable fraction has bounds  $[-\infty, 1]$ .

The estimates of attributable fraction produced by aflogit are identical to those calculated by Benichou and the standard errors are very similar (within 3%).

#### Saved results

aflogit saves the following results in the S\_ macros:

- S\_1 The summary attributable fraction estimate
- S\_2 SE of the summary attributable fraction estimate
- S\_3 Lower confidence limit of the summary attributable fraction
- S\_4 Upper confidence limit of the summary attributable fraction

# Acknowledgments

I am grateful to Tony Swan for pointing the way to estimating attributable fractions from logistic regression and to Heather Dickinson for useful discussions which led to the inclusion of the Poisson regression case.

#### References

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Greenland, S. and K. Drescher. 1993. Maximum likelihood estimation of the attributable fraction from logistic models. Biometrics 49: 865-872.

Tuyns, A. J., G. Pequignot, and O. M. Jensen. 1977. Le cancer de l'oesophage en Ille-et Vilaine en fonction des niveaux de consommation d'alcool et de tabac. Bulletin of Cancer 64: 45-60.

sbe22	Cumulative meta-analysis

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Meta-analysis is used to combine the results of several studies, and the Stata command meta (Sharp and Sterne 1997 and *sbe16.1* in this issue) can be used to perform meta-analyses and graph the results. In cumulative meta-analysis (Lau et al. 1992), the pooled estimate of the treatment effect is updated each time the results of a new study are published. This makes it possible to track the accumulation of evidence on the effect of a particular treatment.

The command metacum performs cumulative meta-analysis (using fixed- or random-effects models) and, optionally, graphs the results.

## Syntax

metacum {theta | exp(theta)} {se\_theta | var\_theta | ll ul [cl]} [if exp] [in range]
 [, var ci effect(f|r) eform level(#) id(strvar) graph
 cline fmult(#) csize(#) ltrunc(#) rtrunc(#) graph\_options]

In common with the commands meta (Sharp and Sterne 1998) and metabias (Steichen 1988), the user provides the effect estimate as *theta* (i.e., a log risk ratio, log odds ratio, or other measure of effect). Likewise, the user supplies a measure of *theta*'s variability (i.e., its standard error, *se\_theta*, or its variance, *var\_theta*). Alternatively, the user provides exp(theta) (e.g. a risk ratio or odds ratio) and its confidence interval, (ll, ul).

#### **Required input variables**

These are the same as for the new version of meta described in sbe16.1 in this issue.

### Options for displaying results

- var means the user has specified a variable containing the variance of the effect estimate. If this option is not included, the command assumes the standard error has been specified.
- ci means the user has specified the lower and upper confidence limits of the effect estimate, which is assumed to be on the ratio scale (e.g. odds ratio or risk ratio).
- effect (f|r) must be included. This specifies whether fixed (f) or random (r) effects estimates are to be used in the output and graph.
- eform requests that the output be exponentiated. This is useful for effect measures such as log odds ratios which are derived from generalized linear models. If the eform and graph options are used, then the graph output is exponentiated, with a log scale for the x-axis.
- level (#) specifies the confidence level, in percent, for confidence intervals. The default is level (95) or as set by set level.
- id(*strvar*) is a character variable which is used to label the studies. If the data contains a labeled numeric variable, then the decode command can be used to create a character variable.

#### **Options for graphing results**

graph requests a graph.

cline asks that a vertical dotted line be drawn through the combined estimate.

- fmult(#) is a number greater than zero which can be used to scale the font size for the study labels. The font size is automatically reduced if the maximum label length is greater than 8, or the number of studies is greater than 20. However it may be possible to increase it somewhat over the default size.
- csize(#) gives the size of the circles used in the graph (default 180).
- ltrunc(#) truncates the left side of the graph at the number #. This is used to truncate very wide confidence intervals. However # must be less than each of the individual study estimates.
- rtrunc(#) truncates the right side of the graph at #, and must be greater than each of the individual study estimates.

graph\_options are any options allowed with graph, twoway other than ylabel(), symbol(), xlog, ytick, and gap.

# Background

The command metacum provides an alternative means of presenting the results of a meta-analysis, where instead of the individual study effects and combined estimate, the cumulative evidence up to and including each trial can be printed and/or graphed. The technique was suggested by Lau et al. (1992).

#### Example

The first trial of streptokinase treatment following myocardial infarction was reported in 1959. A further 21 trials were conducted between that time and 1986, when the ISIS-2 multicenter trial (on over 17,000 patients in whom over 1800 deaths were reported) demonstrated conclusively that the treatment reduced the chances of subsequent death.

Lau et al. (1992) pointed out that a meta-analysis of trials performed up to 1977 provided strong evidence that the treatment worked. Despite this, it was another 15 years until the treatment became routinely used.

Dataset strepto.dta contains the results of 22 trials of streptokinase conducted between 1959 and 1986.

. use strepto, clear (Streptokinase after MI)							
. describe							
Contains data fr obs: vars: size:	rom strep 22 7 638	to.dta		Streptokin	ase after	MI	
<ol> <li>trial</li> <li>trialnam</li> <li>year</li> <li>pop1</li> <li>deaths1</li> <li>pop0</li> <li>deaths0</li> </ol>	byte str14 int int int int int	%8.0gTrial number%14sTrial name%8.0gYear of publication%12.0gTreated population%12.0gControl population%12.0gControl population%12.0gControl deaths					
Sorted by: tria	al						
. list trialnam	year pop	1 deaths1 po	p0 deaths0	, noobs			
trialnam Fletcher	year 1959 1963	pop1 12 21	deaths1 1 4	pop0 11 21	deaths0 4 7		
Dewar 1st European	1963	83	4 20	84	15		
Heikinheimo	1971	219	22	207	17		
Italian	1971	164	19	157	18		
2nd European	1971	373	69	357	94		
2nd Frankfurt	1973	102	13	104	29		
1st Australian	1973	264	26	253	32		
NHLBI SMIT	1974	53	7	54	3		
Valere	1975	49	11	42	9		
Frank	1975	55	6	53	6		
UK Collab	1976	302	48	293	52		
Klein	1976	14	4	9	1		
Austrian	1977	352	37	376	65		
Lasierra	1977	13	1	11	3		
N German	1977	249	63	234	51		
Witchitz	1977	32	5	26	5		
2nd Australian	1977	112	25	118	31		
3rd European	1977	156	25	159	50		
ISAM	1986	859	54	882	63		
GISSI-1	1986	5860	628	5852	758		
ISIS-2	1988	8592	791	8595	1029		

Before doing our meta-analysis, we calculate the log odds ratio for each study, and its corresponding variance. We also create a string variable containing the trial name and year of publication:

- . gen logor=log((deaths1/(pop1-deaths1))/((deaths0/(pop0-deaths0))))
- . gen varlogor=(1/deaths1)+(1/(pop1-deaths1))+(1/deaths0)+(1/(pop0-deaths0))
- . gen str4 yc=string(year)
- . gen str21 trnamy=trialnam+" ("+yc+")"
- . meta logor varlogor, var eform graph(f) id(trnamy) xlab(.1,.5,1,2,10) ltr(0.1)
- > rtr(10) cline xline(1) print b2("Odds ratio") fmult(1.5)

Pooled 95% CI Asymptotic No. o: Method   Est Lower Upper z_value p_value studio Fixed   0.774 0.725 0.826 -7.711 0.000 22	es 
Fixed   0.774 0.725 0.826 -7.711 0.000 22	
Fixed 0.774 0.725 0.826 -7.711 0.000 22	 (p=
	(p=
Random   0.782 0.693 0.884 -3.942 0.000	(p=
Test for heterogeneity: Q= 31.498 on 21 degrees of freedom Moment-based estimate of between studies variance = 0.017	
Weights Study 95%	CI
	Upper
Fletcher (1959)   0.67 0.67 0.16 0.01	1.73
Dewar (1963)   1.91 1.85 0.47 0.11	1.94
1st European (1969)   6.80 6.10 1.46 0.69	3.10
Heikinheimo (1971)   8.72 7.61 1.25 0.64	2.42
Italian (1971)   8.18 7.19 1.01 0.51	2.01
2nd European (1971)   31.03 20.39 0.64 0.45	0.90
2nd Frankfurt (1973)   7.35 6.54 0.38 0.18	0.78
1st Australian (1973)   12.75 10.50 0.75 0.44	1.31
NHLBI SMIT (1974)   1.93 1.87 2.59 0.63	10.60
Valere (1975)   3.87 3.63 1.06 0.39	2.88
Frank (1975)   2.67 2.55 0.96 0.29	3.19
UK Collab (1976)   20.77 15.39 0.88 0.57	1.35
Klein (1976)   0.68 0.67 3.20 0.30	34.59
Austrian (1977)   20.49 15.24 0.56 0.36	0.87
Lasierra (1977)   0.65 0.64 0.22 0.02	2.53
N German (1977)   21.59 15.84 1.22 0.80	1.85
Witchitz (1977)   2.06 1.99 0.78 0.20	3.04
2nd Australian (1977)   10.50 8.92 0.81 0.44	1.48
3rd European (1977)   13.02 10.68 0.42 0.24	0.72
ISAM (1986)   27.13 18.63 0.87 0.60	1.27
GISSI-1 (1986)   303.12 49.69 0.81 0.72	0.90
ISIS-2 (1988)   400.58 51.76 0.75 0.68	0.82

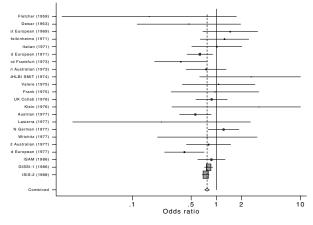


Figure 1: Streptokinase meta-analysis

It can be seen from the fixed-effects weights, and the graphical display, that the results are dominated by the two large trials reported in 1986. We now do a cumulative meta-analysis:

<pre>. metacum logor varlogor, var effect(f) graph eform id(trnamy) xlab(.1,.5,1,2) &gt; ltr(0.1) cline xline(1) b2("Odds ratio") fmult(1.5)</pre>							
Cumulative fixed-effects meta-analysis of 22 studies (exponential form)							
	Cumulative	 95%	CI				
Trial	estimate	Lower	Upper	z	Р	value	
Fletcher (1959)	0.159	0.015	1.732	-1.509		0.131	
Dewar (1963)	0.355	0.105	1.200	-1.667		0.096	
1st European (1969)	0.989	0.522	1.875	-0.034		0.973	
Heikinheimo (1971)	1.106	0.698	1.753	0.430		0.667	
Italian (1971)	1.076	0.734	1.577	0.376		0.707	
2nd European (1971)	0.809	0.624	1.048	-1.607		0.108	
2nd Frankfurt (1973)	0.742	0.581	0.946	-2.403		0.016	
1st Australian (1973)	0.744	0.595	0.929	-2.604		0.009	
NHLBI SMIT (1974)	0.767	0.615	0.955	-2.366		0.018	

	. 1		Oddsi		2
	.1			.5 1	2
ISIS-2 (1988)	-				
ISAM (1986) GISSI-1 (1986)	]				-
d European (1977) ISAM (1986)	1				-
d Australian (1977)	-				-
Witchitz (1977)	4				
Lasierra (1977) N German (1977)	]				_
Austrian (1977)	1				—
Klein (1976)	-				
UK Collab (1976)	4				
Valere (1975) Frank (1975)	1			p	
IHLBI SMIT (1974)	-				
it Australian (1973)	-				_
d European (1971) nd Frankfurt (1973)	]				
Italian (1971) d European (1971)	1				
łeikinheimo (1971)	+		-	•	
st European (1969)	-			•	
Fletcher (1959) Dewar (1963)	1	,	0		
Fletcher (1959)	1 .	_			
1919-7 (1900)	0.114	0.725	0.020	-1.111	0.000
GISSI-I (1986) ISIS-2 (1988)	0.797	0.731	0.870	-5.092 -7.711	0.000
GISSI-1 (1986)	0.784	0.883	0.899	-5.092	0.001
ISAM (1986)	0.784	0.683	0.899	-3.470	0.001
3rd European (1977)	0.010	0.665	0.894	-3.448	0.001
2nd Australian (1977)	0.810	0.695	0.945	-2.688	0.007
Witchitz (1977)	0.810	0.691	0.950	-2.596	0.009
N German (1977)	0.811	0.691	0.951	-2.571	0.010
Lasierra (1977)	0.757	0.637	0.900	-3.150	0.002
Austrian (1977)	0.762	0.641	0.906	-3.072	0.002
Klein (1976)	0.808	0.668	0.976	-2.213	0.027
UK Collab (1976)	0.801	0.662	0.968	-2.296	0.022
Frank (1975)	0.783	0.634	0.968	-2.262	0.024
Valere (1975)	0.778	0.628	0.965	-2.285	0.022

Figure 2: Streptokinase cumulative meta-analysis

By the end of 1977 there was clear evidence that streptokinase treatment prevented death following myocardial infarction. The point estimate of the pooled treatment effect was virtually identical in 1977 (odds ratio=0.771) and after the results of the large trials in 1986 (odds ratio=0.774).

#### Note

The command meta (Sharp and Sterne 1998) should be installed before running metacum.

# Acknowledgment

I thank Stephen Sharp for reviewing the command, Matthias Egger for providing the streptokinase data, and Thomas Steichen for providing the alternative forms of command syntax.

#### References

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-. 1998. sbe16.1: New syntax and output for the meta-analysis command. Stata Technical Bulletin 42: 6-8.

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sbe23	Meta-analysis regression
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The command metareg extends a random effects meta-analysis to estimate the extent to which one or more covariates, with values defined for each study in the analysis, explain heterogeneity in the treatment effects. Such analysis is sometimes termed "meta-regression" (Lau et al. 1998). Examples of such study-level covariates might be average duration of follow-up, some measure of study quality, or, as described in this article, a measure of the geographical location of each study. metareg fits models with two additive components of variance, one representing the variance within units, the other the variance between units, and therefore is applicable both to the meta-analysis situation, where each unit is one study, and to other situations such as multi-center trials, where each unit is one center. Here metareg is explained in the meta-analysis context.

#### Background

Suppose  $y_i$  represents the treatment effect measured in study *i* (*k* independent studies, i = 1, ..., k), such as a log odds ratio or a difference in means,  $v_i$  is the (within-study) variance of  $y_i$ , and  $x_{i1}, ..., x_{ip}$  are measured study-level covariates. A weighted normal errors regression model is

$$Y \sim N(X\beta, V)$$

where  $Y = (y_1, \ldots, y_k)^T$  is the  $k \times 1$  vector of treatment effects, with *i*th element  $y_i$ , X is a  $k \times (p+1)$  design matrix with *i*th row  $(1, x_{i1}, \ldots, x_{ip}), \beta = (\beta_0, \ldots, \beta_p)^T$  is a  $(p+1) \times 1$  vector of parameters, and V is a  $k \times k$  diagonal variance matrix, with *i*th diagonal element  $v_i$ .

The parameters of this model can be estimated in Stata using regress with analytic weights  $w_i = 1/v_i$ . However,  $v_i$  represents the variance of the treatment effect within study *i*, so this model does not take into account any possible residual heterogeneity in the treatment effects *between* studies. One approach to incorporating residual heterogeneity is to include an additive between-study variance component  $\tau^2$ , so the *i*th diagonal element of the variance matrix V becomes  $v_i + \tau^2$ .

The parameters of the model can then be estimated using a weighted regression with weights equal to  $1/v_i + \tau^2$ , but  $\tau^2$  must be explicitly estimated in order to carry out the regression. metareg allows four alternative methods for estimation of  $\tau^2$ , three of them are iterative, while one is noniterative and an extension of the moment estimator proposed for random effects meta-analysis without covariates (DerSimonian and Laird 1986).

#### Method-of-moments estimator

Maximum-likelihood estimates of the  $\beta$  parameters are first obtained by weighted regression assuming  $\hat{\tau}^2 = 0$ , and then a moment estimator of  $\tau^2$  is calculated using the residual sum of squares from the model,

$$RSS = \sum_{i=1}^{k} w_i (y_i - \widehat{y_i})^2$$

as follows:

$$\hat{\tau}_{mm}^2 = \frac{RSS - (k - (p+1))}{\sum_{i=1}^k w_i - \operatorname{tr}(V^{-1}X(X'V^{-1}X)^{-1}X'V^{-1})}$$

where  $\hat{\tau}_{mm}^2 = 0$  if RSS < k - (p+1) (DuMouchel and Harris 1983).

A weighted regression is then carried out with new weights  $w_i^* = 1/\hat{\tau}^2 + v_i$  to provide a new estimate of  $\beta$ . The formula for  $\hat{\tau}_{mm}^2$  in the case of no covariate reduces to the standard moment estimator (DerSimonian and Laird 1986).

## Iterative procedures

Three other methods for estimating  $\tau^2$  have been proposed, and require an iterative procedure.

Starting with  $\hat{\tau}^2 = 0$ , a regression using weights  $w_i^* = 1/v_i$  gives initial estimates of  $\beta$ . The fitted values  $\hat{y}_i$  from this model can then be used in one of three formulas for estimation of  $\tau^2$ , given below:

$$\hat{\tau}_{ml}^{2} = \frac{\sum_{i=1}^{k} w_{i}^{*2} [(y_{i} - \hat{y_{i}})^{2} - v_{i}]}{\sum_{i=1}^{k} w_{i}^{*2}} \text{ maximum likelihood (Pocock et al. 1981)}$$

$$\hat{\tau}_{reml}^{2} = \frac{\sum_{i=1}^{k} w_{i}^{*2} \left[\frac{k}{k - (p+1)} (y_{i} - \hat{y_{i}})^{2} - v_{i}\right]}{\sum_{i=1}^{k} w_{i}^{*2}} \text{ restricted maximum likelihood (Berkey et al. 1995)}$$

$$\hat{\tau}_{eb}^{2} = \frac{\sum_{i=1}^{k} w_{i}^{*} \left[\frac{k}{k - (p+1)} (y_{i} - \hat{y_{i}})^{2} - v_{i}\right]}{\sum_{i=1}^{k} w_{i}^{*}} \text{ empirical Bayes (Berkey et al. 1995)}$$

In each case, if the estimated value  $\hat{\tau}^2$  is negative, it is set to zero.

Using the estimate  $\hat{\tau}^2$ , new weights  $w_i^* = 1/\hat{\tau}^2 + v_i$  (or  $1/v_i$  if  $\hat{\tau}$  is zero) are then calculated, and hence new estimates of  $\beta$ , fitted values  $\hat{y}_i$ , and thence  $\hat{\tau}^2$ . The procedure continues until the difference between successive estimates of  $\tau^2$  is less than

a prespecified number (such as  $10^{-4}$ ). The standard errors of the final estimates of  $\beta$  are calculated forcing the scale parameter to be 1, since the weights are equal to the reciprocal variances.

#### Syntax

metareg has the usual syntax for a regression command, with the additional requirement that the user specify a variable containing either the within-study standard error or variance.

metareg y varlist [if exp] [in range], {wsse(varname) | wsvar(varname) | wsse(varname) wsvar(varname) [ bsest({reml | ml | eb | mm}) toleran(#) level(#) noiter]

The command supplies estimated parameters, standard errors, Z statistics, p values and confidence intervals, in the usual regression output format. The estimated value of  $\tau^2$  is also given.

## Options

- wsse(varname) is a variable in the dataset which contains the within-studies standard error  $\sqrt{v_i}$ . Either this or the wsvar option below (or both) must be specified.
- wsvar(varname) is a variable in the dataset which contains the within-studies variance  $v_i$ . Either this or the wsse option above (or both) must be specified.
- Note: if both the above options are specified, the program will check that the variance is the square of the standard error for each study.
- bsest({reml | ml | eb | mm}) specifies the method for estimating  $\tau^2$ . The default is reml (restricted maximum likelihood), with the alternatives being ml (maximum likelihood), eb (empirical Bayes), and mm (method of moments).
- toleran(#) specifies the difference between values of  $\hat{\tau}^2$  at successive iterations required for convergence. If # is n, the process will not converge until successive values of  $\hat{\tau}^2$  differ by less than  $10^{-n}$ . The default is 4.

level (#) specifies the confidence level, in percent, for confidence intervals. The default is level (95) or as set by set level.

noiter requests that the log of the iterations in the reml, ml, or eb procedures be suppressed from the output.

## Example

BCG is a vaccine widely used to give protection against tuberculosis. Colditz et al. (1994) performed a meta-analysis of all published trials which randomized subjects to either BCG vaccine or placebo, and then had similar surveillance procedures to monitor the outcome, diagnosis of tuberculosis.

The data in bcg.dta are as reported by Berkey et al. (1995). Having read the file into Stata, the log odds ratio of tuberculosis comparing BCG with placebo, and its standard error can be calculated for each study.

. use bcg, clea (BCG and tuberc					
. describe					
Contains data f	rom bcg.d	ta			
obs:	13			BCG and t	uberculosis
vars:	8				
size:	351				
1. trial	str2	%9s		trial ide	entity number
2. lat	byte	%9.0g		absolute	latitude from Equator
3. nt	float	%9.0g		total vac	cinated patients
4. nc	float	%9.0g		total unv	accinated patients
5. rt	int	%9.0g		tuberculo	sis in vaccinated
6. rc	int	%9.0g		tuberculo	osis in unvaccinated
Sorted by:					
. list, noobs					
trial	lat	nt	nc	rt	rc
1	44	123	139	4	11
2	55	306	303	6	29
3	42	231	220	3	11
4	52	13598	12867	62	248
5	13	5069	5808	33	47

6	44	1541	1451	180	372
7	19	2545	629	8	10
8	13	88391	88391	505	499
9	27	7499	7277	29	45
10	42	1716	1665	17	65
11	18	50634	27338	186	141
12	33	2498	2341	5	3
13	33	16913	17854	27	29

. gen logor=log((rt/(nt-rt))/(rc/(nc-rc)))

. gen selogor=sqrt((1/rc)+(1/(nc-rc))+(1/rt)+(1/(nt-rt)))

Note: if either rt or rc were 0, a standard approach would be to add 0.5 to each of rt, rc, nt-rt, and nc-rc for that study (Cox and Snell 1989).

A meta-analysis of the data can now be performed using the meta command described by Sharpe and Sterne (1997 and updated in sbe16.1).

. meta logor selogor, eform graph(r) id(trial) cline xlab(0.5,1,1.5) xline(1)
> boxsh(4) b2("Odds ratio - log scale")

Meta-analysis (exponential form)

1	Pooled	95%	CI	Asymp	totic	No. of
Method				z_value	p_value	studies
Fixed				-10.319	0.000	13
Random	0.474	0.325	0.690	-3.887	0.000	

Test for heterogeneity: Q= 163.165 on 12 degrees of freedom (p= 0.000) Moment estimate of between-studies variance = 0.366

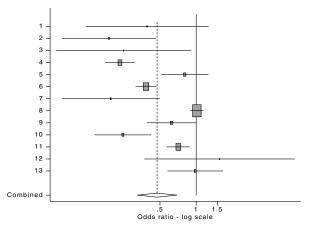


Figure 1: A meta-analysis of the BCG and Tuberculosis data

Both the graph and the statistical test indicate substantial heterogeneity between the trials, with an estimated between-studies variance of 0.366. The random effects combined estimate of 0.474, indicating a strong protective effect of BCG against tuberculosis, should not be reported without some discussion of the possible reasons for the differences between the studies (Thompson 1994).

One possible explanation for the differences in treatment effects could be that the studies were conducted at different latitudes from the equator. Berkey et al. (1995) speculated that absolute latitude, or distance of each study from the equator, may serve as a surrogate for the presence of environmental mycobacteria which provide a certain level of natural immunity against tuberculosis. By sorting on absolute latitude, the graph obtained using meta shows the studies in order of increasing latitude going down the page.

```
. sort lat
. meta logor selogor, eform graph(r) id(trial) cline xlab(0.5,1,1.5) xline(1)
> boxsh(4) b2("Odds ratio - log scale")
(output omitted)
```

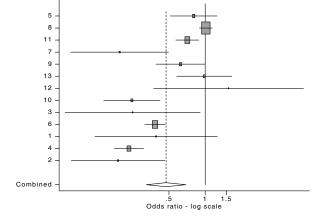


Figure 2: Same as Figure 1 but sorted by latitude

The graph now suggests that BCG vaccination is more effective at higher absolute latitudes. This can be investigated further using the metareg command, with a REML estimate of the between-studies variance  $\tau^2$ .

. metareg logor lat, wsse(selogor) bs(reml) noiter							
Meta-analysis regression					No of studies = 13 tau^2 method reml tau^2 estimate = .0235		
Successive values of tau $^2$ differ by less than $10^{-4}$ - convergence achieved							
		Std. Err.			[95% Conf. Interval]		
lat		.0049432	-6.481 1.977	0.000 0.048	04172470223479 .0029033 .6535356		

This analysis shows that after allowing for additive residual heterogeneity, there is a significant negative association between the log odds ratio and absolute latitude, i.e., the higher the absolute latitude, the lower the odds ratio, and hence the greater the benefit of BCG vaccination. The following plot of log odds ratio against absolute latitude includes the fitted regression line from the model above. The size of the circles in the plot is inversely proportional to the variance of the log odds ratio, so larger circles correspond to larger studies.

```
. gen invvlor=selogor^-2
```

- . gen fit=0.328-0.032\*lat
- . gr logor fit lat [fw=invvlor], s(oi) c(.1) xlab(0,10,20,30,40,50,60)
- > ylab(-1.6094,-0.6931,0,0.6931) l1("Odds ratio (log scale)")
- > b2("Distance from Equator (degrees of latitude)")

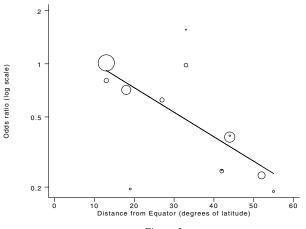


Figure 3

(Note: the axes on this graph have been modified using the STAGE software)

Here a restricted maximum-likelihood method was used to estimate  $\tau^2$ ; the other three methods are used in turn below:

. metareg logor lat, wsse(selogor) bs(ml) noiter

Meta-analys	is regressio	on			No of studies = 13 tau^2 method ml tau^2 estimate = .0037
Successive	values of ta	u^2 differ b	y less tha	un 10 <sup>-4</sup>	- convergence achieved
	Coef.	Std. Err.	Z	P> z	[95% Conf. Interval]
lat   _cons	0327447 .3725098	.0033327 .1043895	-9.825 3.568	0.000	03927670262128 .1679102 .5771093
. metareg l	.ogor lat, ws	se(selogor)	bs(eb) noi	ter	
Meta-analys	is regression	on			No of studies = 13 tau^2 method eb tau^2 estimate = .1373
Successive	values of ta	u^2 differ b	y less tha	un 10 <sup>-4</sup>	- convergence achieved
 	Coef.	Std. Err.	z	P> z	[95% Conf. Interval]
	0305794 .2548214		-3.398 0.812		04822010129388 3603984 .8700413
0	0	(mm) wsse(se erative meth	0		ignored
Meta-analys	is regressio	on			No of studies = 13 tau <sup>2</sup> method mm tau <sup>2</sup> estimate = .0480
	Coef.	Std. Err.	z	P> z	[95% Conf. Interval]
lat   _cons	0315724 .303035	.0061726	-5.115 1.437	0.000 0.151	04367040194744 1102727 .7163427

The estimated value of  $\tau^2$  using a method-of-moments estimator is 0.048, compared with 0.366 before adjusting for latitude, so absolute latitude has explained almost all of the variation between the studies.

The analyses above show that the estimate of the effect of latitude is similar using all four methods. However, the estimated values of  $\tau^2$  differ considerably, with the estimate from the empirical Bayes method being largest. The restricted maximum-likelihood method corrects the bias in the maximum-likelihood estimate of  $\tau^2$ . The basis for using the empirical Bayes method is less clear (Morris 1983), so this method should be used with caution. The moment-based method extends the usual random-effects meta-analysis; below metareg is used to fit a model with no covariate:

. metareg logor, bs(mm) wsse(selogor)							
Meta-analysis regressio	No of studies = 13 tau^2 method mm tau^2 estimate = .3663						
Coef.	Std. Err.						
	.1922628	-3.887	0.000	-1.1242213705641			

Now the estimate of  $\tau^2$  is identical to that obtained earlier from meta, and the constant parameter is the log of the random effects pooled estimate given by meta.

The paper by Thompson and Sharp (1998) contains a fuller discussion both of the differences between the four methods of estimation, and other methods for explaining heterogeneity. Copies are available on request from the author.

## Saved results

metareg saves the following results in the S\_ macros:

- $S_1$  k, number of studies
- S\_2  $\hat{\tau}^2$ , estimate of between-studies variance

## Acknowledgment

I am grateful to Simon Thompson, Ian White, and Jonathan Sterne for their helpful comments on earlier versions of this command.

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sg42.2 Displaying predicted probabilities from probit or logit regression

Mead Over, World Bank, FAX: 202-522-3230

# Syntax

# Description

probpred is an extension of the logpred program of Garrett (1995). It first estimates a probit or logit regression of a dichotomous (or binary) dependent variable on a set of independent variables. The purpose of probpred is to compute and graph the estimated relationship between the predicted probability from this regression and one of the independent variables, holding the others constant. By default, both probpred and logpred display the regression estimates and a graph and listing of the requested predictions and the forecast interval. probpred contains four additional options not included in the original logpred program: logit, one, zero, and level. The default is to estimate a probit regression using the Stata command dprobit, but the logit option instead estimates a logit model using the logistic command. The one and zero options allow the user to specify that some of the covariates listed in option adjust are to be set equal to one or zero instead of to their means. The level option allows forecast intervals to be set to confidence levels determined by the user rather than only to 95% confidence levels.

# Options

from(#) specifies the lowest value of xvar for which a prediction is to be calculated. This option is required.

to (#) specifies the highest value of xvar for which a prediction is to be calculated. This option is required.

inc (#) specifies the increment between adjacent values of xvar. The default increment is 1.

adjust (covlist) specifies the other covariates in the model all of which are set to their sample means in computing the predicted values unless the one or zero options are specified as described below.

one (varlist) specifies a subset of covlist to be set to one instead of to their mean values in the data.

zero(varlist) specifies a subset of covlist to be set to zero instead of to their mean values in the data.

logit specifies that a logit model will be used. The default is probit.

level (#) specifies the confidence level to be used in computing and displaying the forecast interval.

poly(#) indicates that xvar enters the model as a polynomial. Quadratic and cubic models are allowed. They are indicated by poly(2) and poly(3), respectively. The polynomial terms are created and entered in the regression automatically.

nomodel suppresses the display of the estimated regression.

nolist suppresses the list of predicted values.

noplot suppresses the graph of predicted values.

graph\_options allowed include xlabel, ylabel, saving(filename), and titles.

# Example

Using the coronary heart disease data that Garrett used (Garrett 1995), we have that

. probpred chd age, from(20) to(80) inc(5) adj(smk chl)

calculates the predicted probability of coronary heart disease (chd) for five-year increments of age from 20 years to 80 years adjusted for smoking status (smk) and serum cholesterol (chl), and results in the following output:

Iteration Iteration Iteration Probit Es Log Likel	11: Lo 2: Lo 3: Lo timates .ihood =	og Likelihod og Likelihod og Likelihod og Likelihod s = -412.92466	od =-413.3 od =-412.9 od =-412.9	31477 92502 92466		chi2 Prob	er of obs (3) > chi2 do R2	= 51.27 = 0.0000 = 0.0584
chd	1	dF/dx Sto	l. Err.	z	P> z	x-bar	[ 95%	C.I. ]
age smk* chl	.00 10.07	049078 .00 768262 .03 000716 .00	009072 167212 002149	5.32 4.19	0.000	53.7061 .635468	.00313	.006686 .109599
obs. P	.11	165846 )33125 (at						
Probabili Outcome Indepen Covaria Variabl Variabl	ties ar e Varial dent Va tes: .es set .es set bservat	to Zero: to One: tions: 12:	idence Int conary hea e in years a chl 18	cervals art dise s age	ease		eing o	
1.	age 20	pred .0145038	Lowe1 0052657	: 1 7 03!	1pper 52342			
2.	25	.020337	.0088225	5.042	26535			
3.	30	.0280478 .0380526	.0143056	.05	13626			
4.	35	.0380526	.0224407	.06	15699			
5.	40	.0507943 .0667224 .0862665	.0340208	3.073	35753			
6.	45	.0667224	.0497244	1.08	79035			
7.	50	.0862665	.0696591	L .10	56709			
8.	55	.1098053 .1376323	.0925022	2.12	29317			
10.	65 70	.1699221	.1382471	.20	9036			
11. 12.	70	.2066993 .2478149	.1610617	.259	92201			
12. 13.		.2478149			34447			
13.	00	. 232333	.2100358		57741			

The graphical output of the command is given in Figure 1.

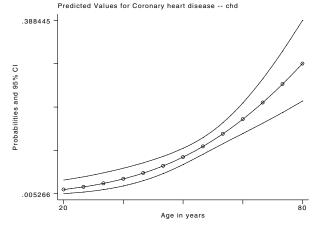


Figure 1. Probability of coronary heart disease as function of age

#### References

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sg76	An approximate likelihood-ratio test for ordinal response models
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# Introduction

An ordinal response is made on a J-category ordered scale. A popular model for  $i = 1, \ldots, n$  ordinal responses  $y_i$  is

$$link(\gamma_{ij}) = \kappa_j - \mathbf{x}_i\beta, \qquad j = 1, \dots, J - 1$$

where  $\gamma_{ij} = \Pr(y_i \leq j | \mathbf{x}_i)$  is the cumulative probability for category j which has associated with it a "cut-point" parameter  $\kappa_j$  related to the cumulative probability of  $y_i$  when  $\mathbf{x}_i = 0$ . The vector  $\mathbf{x}_i$  contains values of explanatory variables and the effects of these variables are quantified by  $\beta$  (a parameter vector of length s). Stata 5.0 has the facility to fit three such models; a model with a logit link can be fitted using the ologit command, a probit link model can be fitted using oprobit, and the complementary-log-log link model can also be fitted because of its equivalence to a continuation-ratio model (Läärä and Matthews 1985) which in turn can be fitted (after appropriate modifications to the data) by specifying a complementary-log-log link for binary data using the glm command.

The model given above defines effects of explanatory variables on cumulative response probabilities that are constant across all categories of the ordinal response. In the case of a logit link this constancy of effect across categories is usually called the proportional-odds assumption, for the complementary-log-log link it is called the proportional-hazards assumption.

Suppose, for example, that  $y_i$  is a measure of smoking frequency with categories "None", "Occasional", and "Frequent", and the model only contains the explanatory variable "Sex", coded as females = 0 and males = 1. Then the difference in effect size between males and females is

$$\operatorname{link}(\gamma_{0j}) - \operatorname{link}(\gamma_{1j}) = \beta_{sex}, \qquad j = 1, 2$$

i.e. a constant effect across the three categories. In terms of underlying prevalence rates, this model specifies that the difference between males and females is that category probabilities are shifted up (if  $\beta_{sex}$  is positive) the response scale for males, relative to females.

In this example we might suspect that the prevalence of "Frequent" smoking is greater for males than females but that the combined prevalence of "Occasional" and "Frequent" smoking is greater for females than males. This implies a sex effect that is not constant across the categories of the response. A generalization of the first model that allows for such a nonconstant effect is

$$link(\gamma_{ij}) = \kappa_j - \mathbf{x}_i \beta_j, \qquad j = 1, \dots, J - 1$$

where now the effect parameters  $\beta_j$  are allowed to vary over the categories.

The purpose of the command omodel is to fit the first model and to test whether the fit to the data provided by the built-in assumption of effect constancy across categories can be improved upon by fitting the one above (in which the effects of all explanatory variables are different at each category).

## Method

The fitting of the first model is described in the Stata manuals under ologit or oprobit as appropriate. The subsequent test performed by omodel is an approximate likelihood-ratio test. Likelihood-ratio tests are described in general by McCullagh and Nelder (1989, Appendix A).

For comparing the first and second models, the likelihood-ratio test is defined as

$$-2\left[l_m(\mathbf{y};\widehat{\beta}_{(1)}^{mle}) - l_m(\mathbf{y};\widehat{\beta}_{(2)}^{mle})\right] \sim \chi^2_{s(J-2)}$$

where  $l_m(.)$  represents the multinomial log-likelihood function (which implicitly takes the specified link function into account) and  $\hat{\beta}_{(k)}^{mle}$  denotes the maximum-likelihood estimate of the parameters in model (k) (here we consider the  $\tau_j$  parameters and the vector  $\beta$  to be included in  $\beta_{(k)}$ ). The problem with using this test is that the second model cannot be fitted in Stata by maximum likelihood. Hence we employ an approximation to the above test using

$$-2\left[l_m(\mathbf{y};\widehat{\beta}_{(1)}^{ib}) - l_m(\mathbf{y};\widehat{\beta}_{(2)}^{ib})\right] \sim \chi^2_{s(J-2)}$$

where  $\hat{\beta}_{(k)}^{ib}$  denote estimates that are obtained from an alternative estimation method which will be referred to as "independent binaries".

In brief, independent binaries estimation of a model for an ordinal response employs the J-1 binary responses defined as

$$y_{ij}^{\dagger} = \begin{cases} 1 & \text{if } y_i \leq j \\ 0 & \text{otherwise} \end{cases}$$

for  $j = 1, \ldots, J - 1$  where  $E(y_{ij}^{\dagger} | \mathbf{x}_i) = \gamma_{ij}$ . The correlation between these binary responses is given by

$$\operatorname{corr}(y_{ij}^{\dagger}, y_{ik}^{\dagger}) = \sqrt{\frac{\gamma_{ij}(1 - \gamma_{ik})}{\gamma_{ik}(1 - \gamma_{ij})}}, \qquad j < k$$

but in estimation this fact is ignored. Hence for estimation purposes, the model is specified in terms of independent binary responses  $y_{ij}^{\dagger}$  and a command for fitting binary responses (e.g. logit or probit) can be used to obtain the estimates.

As with maximum likelihood, the independent binaries method of estimation provides consistent estimates of the model parameters, but the estimation is inefficient compared to maximum likelihood. The approximation  $l_m(\mathbf{y}; \hat{\beta}_{(k)}^{ib})$  will always be smaller than the true likelihood  $l_m(\mathbf{y}; \hat{\beta}_{(k)}^{mle})$  since this quantity is by definition the maximum value of  $l_m(\mathbf{y}; \beta_{(k)})$ . In our experience the approximation is remarkably good, and it certainly should be for large n or any likelihood that is fairly flat around the maximum. Also note that in performing the approximate test, we subtract two quantities that are both underestimates so any error in the approximation will be reduced rather than magnified at this stage.

The null hypothesis of our test is that the (J-1)  $\beta_j$  parameter vectors in the second model are all equal to  $\beta$  as in the first model. The test statistic is the second  $\chi^2$  statistic above and a significant p value is evidence to reject this null hypothesis and accept that the effects of the explanatory variables are not constant across the categories of the ordinal response.

#### Syntax

```
omodel logit varlist [weight] [if exp] [in range]
omodel probit varlist [weight] [if exp] [in range]
```

fweights are allowed and provide a very useful way of dealing with data that are stored in a contingency table format; see the second example below.

# Example of ordinal by ordinal 2-way tables

The data presented by Agresti (1984, 12) on an undesirable side-effect outcome from different operations for treating duodenal ulcer patients are reproduced in Table 1.

m 1 1	1	<b>C</b> 1	1	•	<b>`</b> 1	•	• .	1	· •
Table	1.	( ross-cl	lassificat	10n 01	· dun	nnıng	severity	and	operation
ruore	1.	C1055 C1	abbilleau	ion of	. uun	up mg	Severity	unu	operation

	Dun	Dumping severity				
Operation	None S	Slight	Moderate	Total		
Drainage and vagotomy	61	28	7	96		
25% resection and vagotomy	68	23	13	104		
50% resection and vagotomy	58	40	12	110		
75% resection	53	38	16	107		
Total	240	129	48	417		

Suppose we have these data in Stata as a record for each patient with variables "operate" (numbered 1-4) and "outcome" (response category numbered 1-3). An ordered logit model is fitted to the data using the command

xi: omodel logit outcome i.operate

and the output from the test of the proportional-odds assumption is

chi2(3) = 3.72 Prob > chi2 = 0.2932

If we had data on the patients age and sex we could fit a more elaborate model

xi: omodel logit outcome i.operate\*age i.sex

and of course we could fit an ordered probit model by simply replacing logit with probit in the above commands.

#### Cheese tasting experiment

Suppose that the following data (from McCullagh and Nelder 1989, 175) are to be analyzed with a cumulative logit model to detect for differences in preference of four cheese additives.

Table 2: Multiple-response cheese-tasting experiment

Cheese	$\overline{I^*}$	II	III	IV	V	VI	VII	VIII	IX†	Total
А	0	0	1	7	8	8	19	8	1	52
В	6	9	12	11	7	6	1	0	0	52
С	1	1	6	8	23	7	5	1	0	52
D	0	0	0	1	3	7	14	16	11	52
Total	7	10	19	27	41	28	39	25	12	208
	*I=strong dislike;						†IX=	excel	lent taste	

These data can be entered into Stata most simply as the contingency table of counts that is presented here, i.e., three variables: "response category" (with values 1–9, labeled I–IX), "additive" (values 1–4, labeled A–D) and "count" (which contains the cell counts of the table). When data are stored in this form, omodel can be used as follows

xi: omodel logit response i.additive [fweight=count]

with the result from the approximate likelihood-ratio test statistic in this case being 44.64 on 16 degrees of freedom. Note that the degrees of freedom are less than s(J-2) = 21 because of the zeroes in the table.

The p value for the test is 0.0002 so the null hypothesis is rejected and we conclude that the preferences between the cheeses cannot be described by effects that are constant across all categories.

#### Technical notes

In this second example, the first attempt to use omodel resulted in the error "matsize too small", which is remedied by, e.g. set matsize 150. When omodel performs the approximate test it is necessary to expand the data to (J-1) times n and

Response category

fit two logistic models to the expanded data. If J and n are large, this can take an appreciable amount of time. Also, if as in this example, J is large, then the matsize problem is likely to be encountered.

In the second example, the log-likelihood for the model is -355.67 and the approximation to this from independent binaries estimation is -355.98.

## Saved results

S\_1 Log-likelihood ratio

S\_2 Test degrees of freedom

#### Discussion

The new command omodel provides an approximate likelihood-ratio test for the cumulative logit and probit models. Note that in the case of the complementary-log-log model an exact test can be calculated by fitting the two models with judicious use of glm.

It is to be emphasized that the likelihood-ratio test does not provide a test of whether the first model fits the data adequately. This goodness-of-fit-type interpretation is not the correct conclusion to be reached from a nonsignificant p value. A significant p value provides strong evidence that a more general model is required. A nonsignificant p value shows a lack of evidence that the general model provides an improvement in fit to the data.

The likelihood-ratio tests the first model against a completely general alternative, the second model. For J > 3 there is an intermediate, nested model in which the effect of all explanatory variables is described by two parameters (as opposed to one parameter in the first and J - 1 parameters in the second). This model is defined as

$$link(\gamma_{ij}) = \kappa_j \exp(\mathbf{x}_i \tau) - \mathbf{x}_i \beta, \qquad j = 1, \dots, J - 1$$

where as before, the parameters  $\beta$  describe shifts of the underlying category probabilities up the response scale, and now the parameters  $\tau$  describe concentration of underlying category probabilities towards the middle of the response scale. The above model cannot at present be estimated in Stata but is worthy of consideration since it describes a phenomenon that often occurs with ordinal response scales.

If a significant p value is obtained from the likelihood-ratio test, it is possible that the above model is sufficient to describe all of the non-constancy of effects across categories. A further likelihood-ratio test comparing the second model and the one above would determine whether this was the case. For the data in Example 2, McCullagh and Nelder (1989, 177) examine the deviance reduction due to the above model and report it to be 3.3 on 3 degrees of freedom indicating that the non-proportionality of odds cannot be described in the simple fashion defined by the model above.

In the case of a nonsignificant p value from the likelihood-ratio test, the deviance value can be compared to a  $\chi^2$  distribution on s degrees of freedom. This informal comparison tests whether the above model could provide an improved fit to the data (over the first model) if it accounted for the entire reduction in deviance between the first and second models.

An alternative to the likelihood-ratio test is to test a score-statistic (see McCullagh & Nelder 1989, Appendix A) with a null hypothesis that the general model parameter structure of the second model is equal to the structure defined by the first model. Such a score statistic test is provided by PROC LOGISTIC in SAS (SAS Institute 1989). For the dumping severity example the score statistic is 4.021 on 3 degrees of freedom (p = 0.2592), a similar result to the approximate likelihood-ratio test.

For the cheese tasting example, the test statistic from SAS is 17.29 on 21 degrees of freedom. In contrast to the likelihood-ratio test this is nonsignificant, hence the conclusion from the score test is that there is no evidence to reject the null hypothesis of proportional odds in favor of the general model. Note that the degrees of freedom presented by SAS makes no allowance for the sparse nature of the data.

Note that in the case of a single parameter the approximate theoretical error associated with a score statistic is of order  $n^{-1/2}$  whereas for the likelihood-ratio test the approximate theoretical error is of order  $n^{-1}$ , hence the latter is theoretically preferable. The advantage of the score statistic in practice is in terms of computing effort, with only the first model needing to be fitted in comparison to the likelihood-ratio test which requires the fitting of both the first and second models.

## References

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sg77	Regression analysis with multiplicative heteroscedasticity	
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One of the main assumptions in the standard linear regression model—as estimated in Stata by the commands regress and fit—is that the residuals are homoscedastic, i.e., all residuals have the same variance  $\sigma^2$ . What are the consequences when we use OLS-estimates while the homoscedasticity assumption is violated, i.e., the residuals are heteroscedastic? It is well-known that the OLS-estimates  $\hat{\beta} = (X'X)^{-1}Xy$  are still unbiased and consistent under standard regularity conditions. However, the standard OLS-based standard errors estimated as  $\hat{\sigma}^2(X'X)^{-1}$  are wrong, and hence most inferences about the regression coefficients are in jeopardy. It is possible to use a different estimator of the standard errors of the regression coefficient, known as the Huber/White/robust/sandwich estimator, and available in Stata with the option robust. This estimator of the standard error is consistent even if the residuals are heteroscedastic. Consequently, tests of hypotheses about regression coefficients that are based on these robust standard errors are consistent as well.

Robust estimation of standard errors, however, is *not* a panacea for possible heteroscedasticity. The reason is that under heteroscedasticity the OLS-estimators are *inefficient*: there exist alternative "more accurate" estimators, i.e. estimators with a uniformly smaller variance. This is known as the Gauss–Markov theorem in the theory of the linear model (Eaton 1983, Ch. 4). Consequently, the "robust" test of hypotheses on parameters is suboptimal.

How can we diagnose whether the homoscedasticity assumption is violated in some regression problem? A variety of tests have been proposed. Currently, Stata contains a score test (hettest) of homoscedasticity against a parametric model that specifies how the residual variance depends on a vector of covariates. In a variant of this score test, the residual variance is specified as a function of the expected value of the dependent variable. Stata's reference manual is silent on how to proceed if any of these tests are significant, i.e., if the homoscedasticity hypothesis has to be rejected. Heteroscedasticity of the latter form, i.e.,  $\sigma^2 = f(Ey)$ , can usually be dealt with by a transformation of the dependent variable. The Box-Cox regression model, in which the transformation is estimated within the Box-Cox family, is quite suitable here. However, heteroscedasticity of the first type is more awkward. Here, we usually cannot avoid actually modeling the variance of the residual (or the dependent variable) in terms of covariates, just like we model the expected value of the dependent variable in terms of some covariates.

The command regh estimates a linear regression model with normally distributed residuals with the specific functional form for heteroscedasticity that is also assumed by hettest,

$$y_i = \mu_i + \sigma_i e_i$$
  

$$\mu_i = \mathbf{E} y_i = \beta_0 + \beta_1 x_{i1} + \ldots + \beta_k x_{ik}$$
  

$$\sigma_i^2 = \mathbf{Var} y(i) = \exp(\gamma_0 + \gamma_1 z_{i1} + \ldots + \gamma_m z_{im})$$

where  $y_i$  is a random variable (the "dependent variable") with mean  $\mu_i$  and variance  $\sigma_i^2$ , and  $x_i$  and  $z_i$  are (vectors of) covariates predicting the mean and log-variance of y respectively. Thus, we have a linear model for the expected value (mean), and a log-linear model for the variance of a response variable, conditional on a set of covariates that predict the mean and variance. In addition, the residuals  $e_i$  are assumed to be standard normal distributed and independent. (See the option cluster below for cluster-correlated observations, and for estimating a model with nonnormal residuals.) The (vector-) coefficients  $\beta$  and  $\gamma$  are to be estimated.

# Syntax diagrams

eqmean is an equation that contains the dependent variable, followed by the x-variables. eqlnvar is an equation that contains the z-variables. A constant is automatically appended to both eqmean and eqlnvar. (Thus, if eqlnvar is empty, regh fits the same model as regress.) In the current implementation, the constants cannot be dropped. regh typed without arguments replays results. regh interfaces to post-estimation commands such as test, predict etc. in the standard way.

reghv simply sets up appropriate equations (named lp\_mean and lp\_lnvar), and interfaces to regh. Thus, to replay results type regh.

Note that the current implementations of regh and reghv do not support weights.

# Options

var(varlist) specifies the variables used to model the log(variance) of the residuals. (Only with reghv).

- robust specifies that the robust method of calculating the (co)variance matrix is to be used instead of the traditional calculation (Harvey 1976). The robust variant of the covariance matrix is also computed for the 2SLS estimator.
- cluster(varname) implies robust and specifies a variable on which clustering is to be based. The cluster-variable may be of type string.
- twostage specifies that Harvey's 2SLS estimator (and the associated consistent (co)variance matrix estimate) should be computed, otherwise the maximum-likelihood estimator is used. If the residuals are normally distributed, the variance of maximum-likelihood estimators is approximately half the variance of the 2SLS-estimator.

level(#) specifies the confidence level, in percent, for confidence intervals. The default is level(95) or as set by set level.

- from(matname) specifies a matrix (row-vector) with initial values. from should be properly named (see online help for ml for details). from enables efficient bootstrapping where one may use "full sample" estimates as starting values for the resamples.
- *maximize\_options* control the maximization process; see [R] **maximize**. You should never have to specify the more technical of these options, although we do recommend specifying the trace option.

#### Additional programs

We have a number of additional programs (commands) to test for and model heteroscedasticity in a regression-like context. Contact the author if you are interested in any of these commands.

- htest is a modification of Stata's hettest that easily provides univariate tests as well.
- white is White's test for homoscedasticity  $(H_0)$  against general forms of heteroscedasticity  $(H_a)$  (see White 1980).
- szroeter is Szroeter's semi-parametric test whether the residual variance is monotone in an exogenous discrete or continuous variable.
- regh2 is similar to regh but uses ml/deriv2 (but Fisher-scoring rather than Newton-Raphson) rather than the hand-coded alternating scoring algorithm. regh2 is considerably slower, but, may be more stable for very ill-conditioned problems.
- reghf computes maximum-likelihood estimators for (location, scale) models with nonnormal residuals (e.g., Cauchy, logistic). There is some evidence that inference on variance-models in the context of regression models is less robust than inference on the model of the mean. Using reghf it is possible to verify whether conclusions depend on the assumption of normality. In case of doubt, use the 2SLS-estimator that is less efficient than the maximum-likelihood estimation under normality, but consistent for nonnormal residuals as well.
- reghv computes maximum-likelihood estimators (for normally distributed residuals) for a variant of the multiplicative heteroscedasticity model that assumes

$$\log \sigma_i^2 = \alpha \mu_i + \gamma' z_{i*}$$

where  $\alpha$  is a scalar parameter.

probith estimates probit models with multiplicative heteroscedasticity. Note that the probit model can be interpreted as a regression model with normally distributed residuals and severe missing values in the dependent variable y: it is only observed when  $y \ge 0$ . This generalizes immediately to the heteroscedastic probit regression model.

#### Example

In this stylized example, I discuss an analysis of data on the management of R&D cooperative alliances, modeled after Blumberg (1997). We wanted to analyze the extent to which firms that engage in an R&D-oriented transaction with a partner, invest in ex-ante planning such as contracting (contract). From Williamson's transactions-costs economics and Granovetterlike sociological arguments about the effects of social embeddedness of the functioning of markets, it is expected that these investments increase with the risks involved (here indicated by the variables volume (log-volume-in-money) and depend (mutual dependency)), decrease to the extent that the partners involved have a common past (past) and expect to have a common future (future), and decrease to the extent that the partners have common ties to other firms (network).

. fit contract volume depend past future network

Source	SS	df	MS		Number of obs F( 5, 88)	
Model	458.319345	5 91.	663869		Prob > F	= 0.0000
Residual	161.770291	88 1.83	829876		R-squared	= 0.7391
+					Adj R-squared	= 0.7243
Total	620.089636	93 6.66	763049		Root MSE	= 1.3558
contract	Coef.	Std. Err.	t	P> t	[95% Conf.	Interval]
+						
volume	.9070136	.1587244	5.714	0.000	.5915822	1.222445
depend	.4112368	.1490966	2.758	0.007	.1149387	.7075349
past	224738	.0171927	-13.072	0.000	2589049	190571
future	218904	.1034922	-2.115	0.037	4245731	013235
network	3401932	.1011277	-3.364	0.001	5411632	1392232
_cons	-1.679443	1.160729	-1.447	0.151	-3.986149	.6272628

Let us compare these results with OLS-estimates with "robust" standard errors.

. fit contract volume depend past future network, robust

Regression	with robust	standard er	rrors		Number of obs F( 5, 88) Prob > F R-squared Root MSE	
contract	Coef.	Robust Std. Err.	t	P> t	[95% Conf.	Interval]
volume	.9070136	.1904846	4.762	0.000	.5284655	1.285562
depend	.4112368	.1038016	3.962	0.000	.2049529	.6175207
past	224738	.0259251	-8.669	0.000	2762586	1732174
future	218904	.0827287	-2.646	0.010	3833099	0544982
network	3401932	.0987157	-3.446	0.001	53637	1440164
_cons	-1.679443	.898974	-1.868	0.065	-3.465965	.1070788

The changes in estimates for standard errors are generally conservative. Next we test for homoscedasticity with a Weisberg-Cook's score test using htest.

. htest,	rhs u			
Score tes	ts for heteroscedasticity	(Ho:	Constant	variance)

1	var = f	(a +	b*z)	
Variable	score	df	р	
+				-+
volume	40.22	1	0.0000	
depend	0.09	1	0.7604	
past	94.68	1	0.0000	
future	2.42	1	0.1201	
network	0.27	1	0.6021	
+				-+
combined	128.14	5	0.0000	T

According to the univariate tests of htest, it appears that the variance of the residuals can be modeled using only volume and past. White's omnibus test for heteroscedasticity also indicates that the regression analysis is misspecified, possibly due to heteroscedasticity.

The multiple-equations command regh expects input via equations.

```
. eq mean : contract volume depend past future network . eq var : volume past
```

. regh mean	var						
iteration 1:       Log-likelihood = -80.76063        delta-b        2.126929         iteration 2:       Log-likelihood = -78.22403        delta-b        .2676002         iteration 3:       Log-likelihood = -78.05332        delta-b        .0455286         iteration 4:       Log-likelihood = -78.04507        delta-b        .0167279         iteration 5:       Log-likelihood = -78.04482        delta-b        .0032507         iteration 6:       Log-likelihood = -78.04482        delta-b        .0005627         Multiplicative heteroscedastic regression       Number of obs = 94         Estimator: mle       Model chi2(7) = 288.008							
Log Likelih	ood	= -78	3.045		Prob ≻ chi2 Pseudo R2 VWLS R2		
contract	Coef.	Std. Err.	z	P> z	[95% Conf.	Interval]	
	.631765 .4844798 2116046 4432523	.0351336 .0297727 .005701 .0231963	17.982 16.273 -37.117 -19.109	0.000 0.000 0.000 0.000	.5629045 .4261265 2227784 4887163	.7006255 .5428332 2004308 3977883	
network   _cons	2814385 .2489145	.0231456	-12.159 0.973		326803 2527041		
var   volume   past   _cons	.5516058 2210131 -3.527318	.1561316 .0179103 .9874938	3.533 -12.340 -3.572	0.000 0.000 0.000	.2455935 2561166 -5.462771	.857618 1859097 -1.591866	

The multi-panel output will look familiar to experienced Stata users. According to the estimated model, (the expected amount of) contracting increases with the volume and dependency associated with the transaction, and decreases with common past, expected common future, and common network ties. The variance of the residuals increases with the volume of the transaction, and decreases with a common past. Comparing the output of fit, robust to regh, it is obvious that the robust standard errors of OLS-estimates are much larger than those of the regh-model, and so the 95% confidence intervals have shrunk drastically.

Having improved our estimates of regression-coefficients, we cannot, however, be very confident that we modeled the heteroscedasticity correctly. Thus, we may be estimating the standard errors of the regh-coefficients inconsistently. This problem can be remedied via robust estimation of standard errors in the regh model.

. regh mean var, robust nolog							
Multiplica Estimator:	tive heteros mle		Number of obs Model chi2(7) Prob > chi2	= 288.008			
Log Likelihood = -78.045					Pseudo R2 VWLS R2		
		Robust					
contract		Std. Err.			[95% Conf.	Interval]	
me an							
volume	.631765	.0373337	16.922	0.000	.5585923	.7049378	
depend	.4844798	.0301037	16.094	0.000	.4254776	.5434821	
past	2116046	.0057706	-36.669	0.000	2229148	2002944	
future	4432523	.024733	-17.922	0.000	4917281	3947766	
network	2814385	.0164501	-17.109	0.000	3136801	2491969	
_cons	.2489145	.2852164	0.873	0.383	3100994	.8079285	
var							
volume	.5516058	.1722091	3.203	0.001	.2140822	.8891293	
past	2210131	.0208703	-10.590	0.000	2619182	1801081	
_cons	-3.527318	1.090857	-3.234	0.001	-5.665359	-1.389277	

Comparing the standard errors of regh with the robust standard errors of regh, robust, we conclude the effects of the possible misspecification of heteroscedasticity are apparently rather mild.

Note that regh produces a trace of the iterative process of computing maximum-likelihood estimates that is similar to the standard Stata's trace—I added the L2-norm of the change in the parameter vectors to better trace progress for ill-conditioned problems for which convergence was problematic before I introduced step-halving in the algorithm.

So far we have shown output of **regh** for maximum-likelihood estimators based on the assumption that the residuals are normally distributed. If you distrust this assumption strongly, you may prefer Harvey's two-stage least squares estimates (2SLS). Note that under normality, the approximate standard errors of maximum-likelihood estimators are roughly 1.4 times smaller than those of 2SLS estimators (Harvey 1976).

. regh mean var, twostage							
Multiplicative heteroscedastic regression Estimator: 2sls					Number of obs Model chi2(7) Prob > chi2	= 222.900 = 0.000	
Log Likelihood		= -110.599			Pseudo R2 VWLS R2	= 0.5019 = 0.9027	
contract	Coef.	Std. Err.	z	P> z	[95% Conf.	Interval]	
me an							
volume	.9070136	.0651181	13.929	0.000	.7793844	1.034643	
depend	.4112368	.0543363	7.568	0.000	.3047396	.517734	
past	224738	.0088016	-25.534	0.000	2419888	2074872	
future	218904	.0426059	-5.138	0.000	30241	1353981	
network	3401932	.0420307	-8.094	0.000	4225718	2578146	
_cons	-1.679443	.4606367	-3.646	0.000	-2.582274	7766116	
var							
volume	.3422644	.2452509	1.396	0.163	1384185	.8229472	
past	1401075	.0281334	-4.980	0.000	1952479	0849671	
_cons	-2.081192	1.551151	-1.342	0.180	-5.121393	.9590086	

# Notes and references

Harvey (1976) suggests a 2SLS-estimator in which  $y_i$  is regressed on  $x_i$  to estimate  $\beta$ , and to regress the log-squared-residuals of this regression on  $z_i$  to estimate  $\gamma$ . After adding 1.2704 to  $\hat{\gamma}_0$ , this estimator of  $\beta, \gamma$  is consistent (Harvey 1976). This estimator is available with the option twostage.

To compute maximum-likelihood estimators for normally distributed residuals, regh uses Harvey's alternating scoring algorithm, starting from the 2SLS estimator (See also Greene 1993). I added step-halving to improve stability of the algorithm. In my experience, this algorithm is fast and converges well.

Robust standard errors and the adaptation of the standard errors for clustered observations follows the method implemented by the \_robust command in Stata.

## Acknowledgment

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sg78	Simple and multiple correspondence analysis in Stata

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Correspondence analysis is a technique that often turns out to be helpful when one wants to analyze large cross-tabular data (see Benzecri and Benzecri 1980 or Greenacre and Blasius 1994). It results in a simple graphical display which permits rapid interpretation and understanding of the data. It is currently available in a number of statistical packages (e.g. SPSS, SAS, or BMDP), but not in Stata 5.0. This insert aims at filling this gap (at least partly). The commands coranal and mca provide the general features of simple and multiple correspondence analysis and are designed for standard applications. Admittedly, they might be inefficient for more specific uses and suffer from some deficiencies outlined below. As I am not myself an experienced

user of these techniques, I will neither enter into technicalities nor provide precise explanations on how to use and interpret the results. Furthermore, the very simple example provided here is just intended to give an idea about how the output can look; it does not substitute for an introductory text. The references given at the end of the insert should accomplish these tasks much better than I can.

The two major drawbacks of the current versions of the procedures are 1) the aspect ratio distortion of the maps (the scales on the vertical axes are usually different from that on the horizontal axes—this is innocuous in most graphs but it may sometimes be misleading if one is not aware of this); it may be necessary to edit the graphs to solve this problem, and 2) the absence of an option for including supplementary points in the analysis; this ought to be made available in the future.

Future developments of these commands should also include the creation of immediate command versions or quasi-immediate commands; that is, using cross-tabulations or matrices as inputs and not just raw data. However, a trick for doing so with the current version is suggested in the help file for coranal.

## Simple correspondence analysis

The command coranal produces numerical results as well as graphical outputs for simple correspondence analyses (that is, analyses of two-way cross-tabulations of variables, or said differently, analyses of contingency tables). The computation algorithm draws, on an almost step-by-step basis, on Blasius and Greenacre (1994).

# Syntax

 $\texttt{coranal } varl \ var2 \ [weight] \ [\texttt{if } exp] \ [\texttt{in } range] \ [\texttt{, } \texttt{d}(\texttt{\#}) \ \texttt{q}(\texttt{\#}) \ \underline{\texttt{asymmetric}}]$ 

aweights, fweights, and iweights are allowed.

# Options

- d(#) specifies the number of dimensions to be considered (for both numerical and graphical displays). If d(0) is specified, then coranal provides no graphical display and returns the numerical output for all nontrivial dimensions. For maps to be readable, # must be set larger than 1. Furthermore, consistent maps can only be obtained by specifying # lower than or equal to the number of underlying non-trivial dimensions. The default # is 0.
- q(#) specifies a quality of representation threshold ( $0 < \# \le 1$ ). It restricts the mappings to points satisfying the condition that their quality of representation (sum of contributions of principal axes) in the d(#) first dimensions is higher than or equal to #. Rejected points are still mapped but symbolized by a dot.
- asymmetric specifies that the joint displays of *var1* and *var2* are to be presented in the form of asymmetric maps (both variables are taken as vertices consecutively). By default, symmetric maps are displayed.

# Example

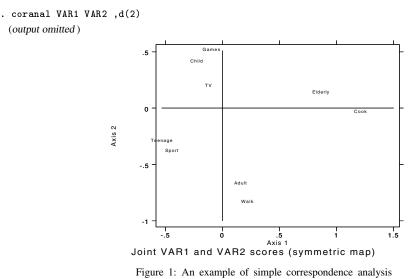
To briefly illustrate the command, consider the following example. The dataset exca.dta (included with this insert) contains a fictional survey. Available individual characteristics are the main leisure-time activity (VAR1), the age group (VAR2), the place of birth (VAR3) and the gender (VAR4). Say we want to explore the links between age groups and leisure-time activity.

. tab VAR1 V	VAR2 ,chi row				
	VAR2				
VAR 1	Child	Teenage	Adult	Elderly	Total
TV			3	4	24
	50.00	20.83		16.67	100.00
Walk	+   1	1		1	7
	14.29	14.29		14.29	
Cook		0			
	11.11 +	0.00	22.22		100.00
Sport				1	18
-	27.78	44.44		5.56   +	
Games	+				17
		11.76		17.65	100.00
Total	+   30			т	75
	40.00	21.33	18.67	20.00	100.00
Pea	arson chi2(12)	= 33.9076	Pr = 0.0	01	

This table and the  $\chi^2$  statistic reveals that there indeed exists a strong relation between the two variables. Let us apply now a simple correspondence analysis on this table for an easy representation of the apparent associations between the various categories (see the references given below for complete explanations about the interpretations of the following numerical and graphical output).

		SIMPLE	CORRESPON	DENCE ANA	ALYSIS
Totol	Inertia :	0.4			
	ipal Inert				
	-		•	•	
	Inertia 0.235		Cumul 0.519		
Dim2	0.173		0.902		
Dim3	0.045	0.098	1.000		
VAR1	coordinate	s:			
	Mass	Inertia	Dim1	Dim2	Dim3
VAR1:1	0.320	0.016	-0.121	0.191	0.014
	0.093				
VAR1:3 VAR1:4	0.120	0.182	1.213 -0.443	-0.039	-0.213
VAR1:4 VAR1:5			-0.443		-0.229 0.133
	coordinate				01100
1 1112			D.≟ 4	D÷0	D.÷
VAR2:1		1nert1a 0 092	Dim1 -0.224	Dim2 0.406	
VAR2:3	0.213 0.187	0.103	0.159	-0.673	0.269
VAR2:4	0.200	0.160	0.866	0.132	-0.177
Expla	ined inert	ia of axi	s by VAR1	:	
	Dim1	Dim2	Dim3		
	0.0198				
	0.0193				
	0.7518 0.2006				
	0.0085				
Expla	ined inert	ia of axi	s by VAR2	:	
-	Dim1		-		
VAR2:1	0.0856				
VAR2:2	0.2558	0.1086 0	.4222		
	0.0202		.3040		
	0.6384		.1415		
Contr	ibutions o			VAR1 :	
VAD1.1	Dim1				
	0.2850 0.0488				
	0.9690				
VAR1:4			.1310		
VAR1:5	0.0307	0.9071 0	.0623		
Contr	ibutions o	f princip	al axes to	VAR2 :	
	Dim1	Dim2	Dim3		
VAR2:1	0.2185	0.7175 0	.0641		
VAR2:2			.1925		
VAR2:3			.1316		
VAR2:4 var11b		0.0219 0	.0395		
	1 T	'V			
		alk			
		ook			
		port ames			
var21b					
vai 210		hild			
		eenage			
		dult			
	4 E	lderly			

The first table reveals that it is possible to obtain a two-dimensional graphical representation of the observed relation between VAR1 and VAR2 by, very roughly speaking, retaining 90 percent of information as Dim1 and Dim2 account for 90 percent of total inertia (note that the total inertia is equal to the  $\chi^2$  statistic divided by the number of observations).



On this graph, we can now observe how (dis-)similar are the profiles of the row points (or column points): categories that are similar to each other regarding their distribution over the categories of the other variable are plotted close to each other in the figure. Basically, the distances between the row points (or column points) in this display are to be seen as the best approximations in two dimensions of the  $\chi^2$  distances between the row profiles (or column profiles) in Euclidian space. Note that the exact location of a point is situated right in the middle of the label name.

By default, coranal produces such "symmetric" maps which are usually more readable. Yet the distances between VAR1 points and VAR2 points are not directly comparable. To interpret such distances one has to display "asymmetric" maps (set the asymmetric option). We might also seek to avoid interpreting points for which the quality of representation is judged to be low (below 85 percent, say), where quality is measured by the contributions of the first two principal axes. Note that this is a very high threshold level.

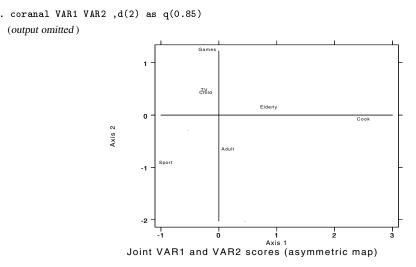


Figure 2: Using the asymmetric and q options

The points representing the profiles of the "Teenage" category and the "Walk" activity have been removed from the display since their quality of representation is below 85 percent. In this display, the activities have been taken as vertices to which the position of the age groups are to be compared. We could now observe, for instance how "Cook" is related to the remaining age groups: this occupation is apparently more a delight for the "Elderly" then for the "Adult" and eventually for the "Child" since the distance between the vertex "Cook" and the "Elderly" group is the smallest, the next smallest being with the "Adult" group and so on. This actually represents the row profile of "Cook" in the cross-tabulation of VAR1 and VAR2 above in the right order.

#### Multiple correspondence analysis

The command mca produces numerical results and graphical representations for multiple correspondence analyses (MCA).

It must be emphasized that there is some controversy about the use and implementation of multiple correspondence analyses (see Greenacre 1984, 1993, 1994). mca actually conducts an adjusted simple correspondence analysis on the Burt matrix constructed with *varlist* (i.e. matrix of frequency counts resulting from all two-way cross-tabulations of the variables in *varlist* including the cross-tabulations of each variable with itself). It can be shown that the total inertia of the Burt matrix is high due to the fitting of the diagonal sub-matrices. Consequently, a simple correspondence analysis applied to this matrix usually results in maps of apparently poor quality. As a remedy, if not otherwise specified, mca adjusts the obtained principal inertias (eigenvalues) following a method suggested by Benzecri and presented in Greenacre (1984).

The computation algorithm draws most largely on Blasius and Greenacre (1994) and Greenacre (1994).

#### Syntax

mca varlist [weight] [if exp] [in range] [, d(#) q(#) <u>n</u>otrans]

aweights, fweights, and iweights are allowed.

# Options

- d(#) specifies the number of dimensions to be considered (for both numerical and graphical displays). If d(0) is specified, then mca provides no graphical display and returns the numerical output for all nontrivial dimensions. For maps to be readable,
   # must be set larger than 1. Furthermore, consistent maps can only be obtained by specifying # lower than or equal to the number of underlying nontrivial dimensions. Default # is 0.
- q(#) specifies a quality of representation threshold (0 <#≤ 1). It restricts the mappings to points satisfying the condition that their quality of representation (sum of contributions of principal axes) in the d(#) first dimensions is higher than or equal to #. Rejected points are still mapped but symbolized by a dot. Note that the appropriateness of this option is somewhat questionable in the case of multiple correspondence analyses due to the above outlined inertia inflation of Burt matrices.</p>

notrans requests that no adjustments to eigenvalues be made.

# Example

To illustrate mca, let us return to our fictional survey dataset. We want now to have an idea about how all the variables relate to each other and not only VAR1 and VAR2. Say we conjecture that keeping two dimensions will retain enough information. We can use

```
. mca VAR1 VAR2 VAR3 VAR4, d(2)
                  MULTIPLE CORRESPONDENCE ANALYSIS
_____
 Total Inertia :
                     0.108
 Principal Inertia Components :
     Inertia
                Share
                         Cumul
Dim1
       0.060
                0.557
                         0.557
Dim2
       0.027
                0.248
                         0.805
 Coordinates :
          Mass
                Inertia
                           Dim1
                                    Dim2
VAR1_1
                  0.004
                          -0.105
                                   0.155
         0.080
VAR1 2
         0.023
                  0.008
                          0.402
                                  -0.262
VAR1_3
         0.030
                  0.016
                          0.652
                                  -0.037
VAR1_4
         0.060
                  0.011
                          -0.309
                                  -0.300
VAR1_5
         0.057
                  0.004
                          -0.034
                                   0.226
         0.100
                  0.006
VAR2_1
                          -0.113
                                   0.195
VAR2 2
         0.053
                  0.011
                                  -0.251
                          -0.371
VAR2_3
         0.047
                  0.011
                          0.338
                                  -0.243
VAR2_4
         0.050
                  0.009
                          0.306
                                   0.105
VAR3 1
         0.103
                  0.005
                          0.192
                                   -0 087
VAR3_2
         0.027
                  0.007
                          0.002
                                   0.369
VAR3_3
         0.120
                  0.005
                          -0.165
                                   -0.007
VAR4 1
         0.110
                  0.007
                          -0.239
                                  -0.008
VAR4 2
         0.140
                  0.005
                          0.188
                                   0.006
 Explained inertia of axes
                          :
                 Dim2
         Dim1
VAR1_1 0.0148 0.0717
VAR1 2 0.0629 0.0599
```

VAR1_3	0.2120	0.0015							
VAR1_4	0.0954	0.2016							
VAR1_5	0.0011	0.1084							
VAR2_1	0.0211	0.1421							
VAR2_2	0.1221	0.1261							
VAR2_3	0.0887	0.1027							
VAR2_4	0.0778	0.0205							
VAR3_1	0.0631	0.0293							
VAR3_2	0.0000	0.1355							
VAR3_3	0.0546	0.0002							
VAR4_1	0.1044	0.0003							
VAR4_2	0.0820	0.0002							
Contri		of princ	ipal	axes	:				
	Dim1	Dim2							
VAR1_1	0.2389	0.5144							
VAR1_2	0.4695	0.1991							
VAR1_3	0.7990	0.0025							
VAR1_4	0.5110	0.4807							
VAR1_5	0.0167	0.7186							
VAR2_1	0.2229	0.6668							
VAR2_2 VAR2_3	0.6621	0.3043 0.2513							
VAR2_3	0.4874	0.2513							
VAR3_1	0.7664	0.1585							
VAR3_1 VAR3_2	0.0004	0.5529							
VAR3_3	0.6792	0.0012							
VAR4_1	0.9385	0.0011							
VAR4_2	0.9385	0.0011							
						1		1	1
			.4 -		VA	F 3_2			$\neg$
						_			
			.2 -		VAR VAR2_1	1_5			F
					VAR1_1				
						VAR	2_4		
		2				VAR4_2			
		Axis	0 -		VAR¥AR3_3			VAR1_3	
						VAR3_1			
			2 -						ŀ
				VAR2_	_2 .R1_4	VA VA	R2_3 VAR1_2		
				VA	un i_4				
			4 -						L
				5		0		5	
						Axis 1			
				Mult	iple Corre	espondenc	e Ana	lysis	
						Figure 3			

Figure 3

Unfortunately, maps resulting from multiple correspondence analyses may result in rather inextricable displays (as in this example) which may be difficult to interpret. By looking carefully at the numerical output and the figure, we can observe that the preceding MCA reveals, for instance, a certain tendency for the activities of the type "Walk" (VAR1\_2) and "Cook" (VAR1\_3) to be more associated with women (VAR4\_2) while "TV" (VAR1\_1) and "Sport" (VAR1\_4) seem more associated with men (VAR4\_1). It appears also that there is apparently relatively more women in the oldest age groups (VAR2\_3 and VAR2\_4) than in the youngest (VAR2\_1 and VAR2\_2).

#### References

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sg79	Generalized additive models
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[Editor's note: the gam program is only available for Stata for Windows.]

## Introduction

Generalized Additive Models (GAMs) are flexible extensions of Generalized Linear Models (GLMs). Whereas the link function in a GLM is a linear combination of the predictors, in a GAM the link function may include flexible functions of arbitrary complexity based (for example) on smoothing splines. The amount of smoothing applied to each predictor is controlled by the user according to a quantity known as the equivalent degrees of freedom (df). The GLM is the special case with 1 df for each predictor.

GAMs may be used for exploratory analysis when one knows very little about the functional forms in a dataset. Plots of the smooth functions and a related significance test may assist the user to detect nonlinear relationships that were previously hidden. GAMs may also be useful to suggest functional forms for parametric modeling or for checking an existing parametric model for bias. A plot of the smooth function together with the corresponding parametric function for a given predictor may reveal "hot spots" of poor fit.

Generalized additive modeling is implemented in a FORTRAN program named GAMFIT. It was written by T. J. Hastie and R. J. Tibshirani, who introduced and developed the GAM, and is available from Statlib http://lib.stat.cmu.edu/general/. Here we provide an ado-file, gam, which is an interface between Stata and a slightly modified version of the FORTRAN program. We have only implemented the procedure to work on Windows platforms. The user is completely shielded from the original program. gam has the following basic syntax:

gam yvar xvars [if exp] [in range] [major\_options minor\_options]

Full details are given in the section Syntax of the gam command.

In this implementation cubic smoothing splines are used to estimate the flexible functions, although many other types of estimator could have been used such as kernel smoothers or locally weighted running line smoothers (lowess). In addition, gam can fit additive proportional hazards models to survival data. Details of all these methods can be found in Hastie and Tibshirani (1990a).

#### The generalized additive model

We now discuss the generalized additive model, dealing with the additive proportional hazards model in the next section. The link function  $g(\mu)$  for a GAM with q predictors  $x_1, \ldots, x_q$  has the form

$$g(\mu) = \alpha + \sum_{j=1}^{q} f_j(x_j)$$

where the  $f_j$ 's are smooth functions of the predictors. The "additive" in GAM refers to the fact that the predictor effects are assumed to be additive on the scale of the link function. The "generalized" alludes to the fact that we can use the error structures and link functions that are available with generalized linear models.

As a running example, we use data in the file kyph.dat derived from 81 patients who underwent corrective spinal surgery. The aim of the study was to assess the incidence of spinal deformities following surgery, and discover how various predictors affect this. The response y is binary with y = 1 denoting the presence of kyphosis, which is defined to be a forward flexion of the spine of at least 40 degrees, and y = 0 denoting an absence. There are 3 predictors, age at time of surgery (age), the starting range of vertebrae levels involved in the operation (start), and the number of vertebral levels involved (number). These data were collected by Bell et al. (1994) and have previously been analyzed with GAMs by Hastie and Tibshirani (1990a).

We use a logistic additive model for these data and allow each function to have 4 degrees of freedom (df) via the df option. This option is very important since it controls the flexibility of the function estimators. The actual value of df for a predictor may be viewed as the number of parameters it uses. We now illustrate how the value of df can affect function estimates.

We fit the model logit  $(p) = \alpha + f(age)$  using different values of df. We chose the values 1, 4 and 15. Figure 1 shows the fitted values from these models on the probability scale. The 1 df model is the linear model logit  $(p) = \alpha + \beta \cdot age$  and the estimates are the same as those produced by logit or logistic. The estimated smooth functions for the 4 df and 15 df exhibit a large amount of curvature, with the 15 df model displaying implausible wiggles everywhere, showing that it is overfitted. This demonstrates that care needs to be taken when using the df option.

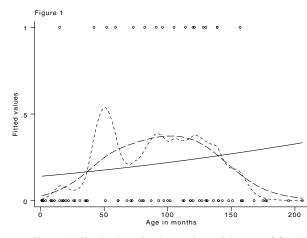


Figure 1: Fitted values for three values of degrees of freedom

The Stata command to fit the multivariable model with age, start, and number is

. gam y age start number, df(4) family(binomial)

which results in the following output:

Generalized	l Additiv	re Model with	family bi	nom, link lo	git.	
Deviance	= 44	3.299 .1354		Dispe	f obs = rsion =	81 1
уl	df		Std. Err.	z	Gain	P>Gain
age	3.874	.0191194	.0055689	3.433	7.929	0.0430
start	4.220	3131638	.0633463	-4.944	7.922	0.0563
number	4.205	.3795253	.1825391	2.079	4.735	0.2159
_cons	1	-2.72886	.311933	-8.748	•	
Total gain	(nonline	arity chisqu	uare) =	20.586 (9.29	9 df), P	= 0.0171

This table is similar in many ways to the table produced using the logit command with columns relating to the linear coefficient, its standard error, and z score. However, there are additional columns relating to the degrees of freedom (df) and gain. The table reveals that the df used with each predictor is not equal to the exact value requested. This is because the program makes its best guess at the requested df initially and calculates the exact values at the convergence of the procedure. These values rarely agree exactly. We have noticed that transforming the predictors to approximate normality (see [R] **Inskew0**) can improve the accuracy of the final values, although the fitted functions will be different.

The gain allows us to assess apparent nonlinearities in the fitted functions. The gain for a given predictor is the deviance increase (i.e. worsening of fit) that occurs if a linear term is used to model the predictor instead of the smooth term. Predictors which are modeled linearly (df = 1) have no entries in the Gain column. The final column refers to an approximate significance test on the smooth function. Under the hypothesis that any nonlinearities in the smooth function are artifactual, the gain is assumed to have a chi-squared distribution with degrees of freedom equal to df - 1, which can be interpreted as the number of degrees of freedom associated with the nonlinear part of the relationship with the outcome variable. We subtract one to allow for the linear component of the smoothing spline fit whose coefficient is given in the Lin. Coef column, followed by its standard error. The nonlinear df is usually noninteger valued and the test statistic has approximately a gamma distribution with the appropriate parameters.

An examination of the table indicates that only the functions for age and start show any signs of nonlinearity. The model may be simplified and we do this using a stepwise procedure suggested by Hastie and Tibshirani (1990a). Each term may be specified as smooth with 4 df, linear, or excluded from the model. Starting with the model with every term specified as smooth we arrive at the model without number, whose command line is

. gam y age start, df(4) family(binomial)

This produces the following table of results:

Generalized	Additiv	e Model with	family bin	nom, link lo	git.	
Model df	=	9.071		No. o	f obs =	81
	= 49			1	rsion =	1
у	df	Lin. Coef.	Std. Err.	z	Gain	P>Gain
•		.0131614	.0062991	2.089	7.826	0.0441
start	4.223	2850923	.0641426	-4.445	9.184	0.0323
_cons		-2.0261	.349529	-5.797	•	•
		arity chisqu				

In addition to the table, we obtain information about the fit from several new variables created automatically by gam. GAM\_mu holds the fitted values on the original scale of the response, and information on the individual predictors is found in the variables prefixed by  $s_{-}$ ,  $e_{-}$  and  $r_{-}$ . For example, including age in the model leads to creation of the variables  $s_{-}age$ ,  $r_{-}age$ , and  $e_{-}age$ . The variables prefixed by  $s_{-}$  are the smooth function estimates, those prefixed by  $e_{-}$  are the pointwise standard errors of the estimates, and those prefixed by  $r_{-}$  are the partial residuals which we now describe in detail.

The algorithm for fitting GAMs involves two iterative loops which are executed until convergence of the estimates occurs. The outer loop, the so-called local scoring algorithm, is similar to the Fisher scoring algorithm used to fit GLMs. This sets up an "adjusted dependent variable" used by the inner loop and recalculates weightings. The inner loop, the backfitting algorithm, fits a weighted additive model with the adjusted dependent variable as the response. The latter is assumed to be normally distributed although the variances are heterogeneous. Within the backfitting algorithm the weighted partial residuals for each predictor are smoothed as functions of the predictor values. The partial residuals for a given predictor are the raw residuals with the effect of the predictor removed. For example, if we denote the adjusted dependent variable by  $z_i$  and the current estimated functions by  $\hat{f}_i(x_{ij})$  then the partial residuals for the *k*th predictor are given by

$$r_{ik} = z_i - \widehat{\alpha} - \sum_{j \neq k} \widehat{f}_j(x_{ij})$$

We give plots of the smooth functions and their pointwise standard errors for age and start in Figure 2. The standard errors give an idea of the accuracy of the estimated function at each observed predictor value but they do not provide global estimates of the uncertainty in the function.

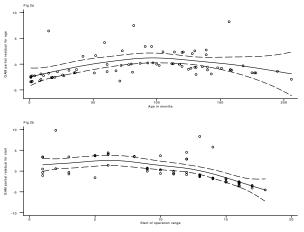


Figure 2: Smooth functions and standard errors for age and start

These plots reveal that each function appears to model the relationships in the data well.

#### The additive proportional hazards model

One can use the gam command to fit the additive proportional hazards model. This model, introduced by Hastie and Tibshirani (1990b), extends the familiar semiparametric model of Cox (1972) by allowing flexible predictor effects. The additive proportional hazards model has the form

$$\lambda(t|x_1,\ldots,x_q) = \lambda_0(t) \exp\left\{\sum_{j=1}^q f_j(x_j)\right\}$$

where  $\lambda_0(t)$  is the baseline hazard and the  $f_j(x_j)$  are smooth functions which are estimated using smoothing splines.

We now present an example of the use of gam with survival data (given in the file pbc.dat) on 312 patients with primary biliary cirrhosis (PBC) who were referred to the Mayo Clinic between 1974 and 1984. They participated in a randomized, placebo-controlled trial of the drug D-penicillamine. Patients were followed up until July 1986 for survival status, and 125 died. The response t is survival time and the predictors we use are age (age), presence of edema (edm), serum bilirubin (bil), albumin (alb) and prothombin time (pro). Of these predictors, edm has 3 categories and is split into two indicator variables edm2 and edm3. The censoring variable is denoted by cens. No difference in the survival times of the treated and untreated patients were detected, so we exclude the treatment variable from the model. More details of the study can be found in Fleming and Harrington (1991).

Initially we fit an additive proportional hazards model using

. gam t age bil alb pro edm2 edm3,dead(cens) df(4, edm2 edm3:1) family(cox)

which allows every predictor 4 df except edm2 and edm3 which are binary and so are given 1 df. This results in the following table:

312 1	of obs = ersion =	No.	family cox,	e Model with 9.606 65.37	= 1	lodel df
P>Gain	Gain	z		Lin. Coef.	df	+- t
0.0666	7.352	3.776	.0090497		4.108	+- age
0.0000	35.737	8.112	.0156951	.1273225	4.687	bil
0.5472	2.546	-5.025	.2099035	-1.054681	4.451	alb
0.2108	5.002	3.076	.0935668	.2878167	4.360	pro
		0.671	.2781	.186678	1	edm2
		3.183	.298307	.949492	1	edm3

Total gain (nonlinearity chisquare) = 50.637 (13.606 df), P = 0.0000

After dropping edm2 and any smooth terms which do not significantly contribute to the fit we arrive at the model presented below:

312 1	of obs = ersion =	Dispe		76.57	= 10	odel df eviance
P>Gain	Gain	z	Std. Err.	Lin. Coef.	df	t
		4.108	.0086492		1	age
0.0000	37.744	8.101	.0158869	.1286926	5.447	bil
		-4.688	.2200582	-1.031641	1	alb
		3.104	.0838921	.2603843	1	pro
		3.315	.285888	.947819	1	edm3

This table reveals that there is a strong nonlinear relationship involving serum bilirubin (bil). Figure 3 shows the smooth function for this variable and suggests that a logarithmic transformation of bil would be suitable as a simple parametric model. No partial residuals are shown as these are not created for the Cox model.

(Figure 3 on the next page)

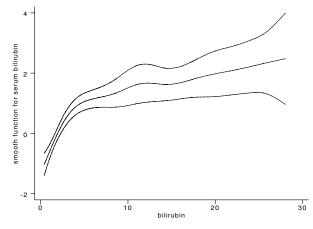


Figure 3: The smooth function for serum bilirubin for PBC data

We note that Figures 2 and 3 were produced using the gamplot program which we also provide. gamplot allows the user to easily plot the smooth functions produced by gam with options to include partial residuals and standard error bands. Full details are given in the section entitled *Syntax of the* gamplot *command*.

## Syntax of the gam command

gam yvar xvars [weight] [if exp] [in range] [, major\_options minor\_options]

aweights and fweights are allowed.

We note that gam expects to find the FORTRAN executables in the directory  $c:\ado$ . If they are not stored here, the global macro GAMDIR needs to be set. For example if the FORTRAN executable is stored in  $d:\mbox{wado}$ , you would enter

global GAMDIR d:\myado\

before calling gam.

## Options

The major\_options (most used options) are

family(family\_name) link(link\_name) df(dflist) dead(deadvar)

where *family\_name* is one of the following

gaussian <u>b</u>inomial poisson gamma <u>c</u>ox

and link\_name is one of the following

<u>ide</u>ntity | <u>l</u>ogit | log | <u>inv</u>erse | <u>c</u>ox

The minor\_options (less used options) are

big <u>mi</u>ssing(#) <u>nocons</u>tant

## **Major options**

family(family\_name) specifies the distribution of yvar. The default is family(gaussian).

- link(link\_name) specifies the link function. The default for each family are the canonical links: identity for family gauss, logit for binomial, log for poisson, inverse for gamma and (by convention) cox for cox.
- df(dflist) sets up the degrees of freedom (df) for each predictor. These values need not be integers. An item in dflist may be either # or xvarlist: #, where xvarlist is a subset of xvars. Items are separated by commas. With the first type of item, the df for all predictors are taken to be #. With the second type of item, all members of xvarlist have # df. If an item of the second type follows one of the first type, the later # overrides the earlier # for each variable in xvars. The default value is 1 for each predictor.

dead(deadvar) only applies to Cox regression where deadvar is the censoring status variable (0 for censored, 1 for "dead").

#### Minor options

big asks for the large-problem version of the GAMFIT FORTRAN program, gambig.exe. The largest problem that can be fit is 70,000 reals in the standard version of GAMFIT (gamfit.exe) and 1,000,000 reals in the big version (gambig.exe). These quantities are the amount of storage needed by the FORTRAN programs, not the amount of data stored in Stata. The problem size is approximated by the following formula:  $reals = 1000 \times N \times (vsize^{0.2})/25$  where N is the number of observations in the problem and vsize is the total number of variables, including the constant and  $cens\_var$  if a Cox model is fit. For example, for a model with a constant and a single predictor (vsize = 2) the biggest problems that can be fit are N = 1523 and N = 21764 for the standard and big versions respectively.

noconstant specifies that the model shall not have an intercept.

missing(#) defines the missing value code seen by gam to be #, which must be a number. The default is 9999. The FORTRAN program is able to impute missing values. See Hastie and Tibshirani (1990a) for more details.

#### Syntax of the gamplot command

gamplot xvar [xvar2] [if exp] [in range] [, noconf nopres se(#) abs(#) graph\_cmd\_options] where xvar is a member of xvarlist. If xvar2 is specified the functions are plotted against this variable.

## Options

noconf suppresses the plotting of the pointwise standard errors.

nopres suppresses the addition of partial residuals on the plot.

- se(#) determines the width of the standard error bands. The default is 0.95 which corresponds to 95% coverage (using assumptions of normality).
- abs (#) is the maximum permitted distance between the smooth function and the partial residuals. Any partial residuals which exceed this value will not be plotted. The user is made aware of this. The default is  $10^{-15}$ .
- graph\_cmd\_options are options appropriate to graph. The only options that cannot be used are sort, symbol, connect, and pen as these are already being used.

#### Warning

We cannot vouch for the results from the FORTRAN software gamfit.exe and have occasionally noticed anomalies. However we believe it to be reliable in the vast majority of instances. gam can fail to converge with Cox regression and can occasionally cause Stata to shut down without warning. We find that this problem can usually be cured by changing the values of *df* slightly.

We also note that (non-binary) predictors are standardized before analysis. As a result the estimate and standard error of the intercept will differ from those produced using Stata commands such as logit, cox, and regress.

#### Acknowledgment

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	sg80	Indirect standardization	
--	------	--------------------------	--

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#### Syntax

istdize casevars popvars stratavars using filename [if exp] [in range],

 $\{popvars(casevar_p popvar_p) \mid rate(ratevar_p \# | crudevar_p)\} [by(groupvars) print <u>format(%fmt) level(#)]</u>$ 

#### Description

istdize produces indirectly standardized rates for a study population based on a standard population. This standardization method is indicated when stratum-specific rates for the population being studied are either unavailable or are based on small samples and thus unreliable. The standardization uses the stratum-specific rates of a standard population to calculate the expected number of cases in the study population(s), sums them, and then compares them to the actual number of cases observed. The standard population's information is contained in another Stata data file and specified via using on the command line.

In addition to the indirectly standardized rate, the command produces point estimates and exact confidence intervals of the study population's standardized mortality ratio (SMR) if death is the event of interest or the standardized incidence ratio (SIR) for studies of disease incidence. Here we refer to both ratios as SMR.

 $casevar_s$  is the variable name for the study population's number of cases (deaths). It must contain integers and each subpopulation identified by *groupvar* must have the same values or missing.

*popvars* identifies the number of subjects represented by each observation in the study population.

stratavars define the strata.

#### Options

popvars( $casevar_p \ popvar_p$ ) or rate( $ratevar_p \ \#|crudevar_p$ ) must be specified. Only one of these two options is allowed. These options are used to describe the standard population's data where:

 $popvars(casevar_p \ popvar_p)$ : casevar\_p records the number of cases (deaths) for each stratum in the standard population. popvar\_p records the total number of individuals in each stratum (individuals at risk).

rate ( $ratevar_p # | crudevar_p$ ):  $ratevar_p$  contains the strata specific rates.  $# | crudevar_p$  is used to specify the crude case rate either via a variable name or optionally by the crude case rate value. If a crude rate variable is used it must be the same for all observations, although, it could be missing for some.

by(groupvars) specifies variables identifying study populations when more than one exist in the data. If this option is not specified the entire study population is treated as one group.

print outputs a table summary of the standard population before displaying the study population results.

format (%fmt) specifies the format to use for displaying the summary table. The default is %10.0g.

level (#) specifies the confidence level, in percent, for the confidence interval of the adjusted rate and the SMR; see [R] level.

#### Remarks

Standardization of rates can be performed via the indirect method whenever the stratum-specific rates are either unknown or unreliable. In situations were the stratum specific rates are known, the direct standardization method is preferred. See [R] **dstdize**.

In order to apply the indirect method the following must be available:

- 1. The observed number, O, of cases in each population to be standardized. For example, if death rates in two states are being standardized using the US death rate for the same time period, then you must know the total number of deaths in each state.
- 2. The distribution,  $n_1, \ldots, n_k$ , across the various strata for the population being studied. If you are standardizing the death rate in the two states adjusting for age, then you must know the number of individuals in each of the k age groups.
- 3. The strata specific rates,  $p_1, \ldots, p_k$ , for the standard population. You must have the US death rate for each stratum (age group).
- 4. The crude rate, C, of the standard population. For example, the mortality rate for all of the US for the year.

Then the expected number of cases (deaths), E, in each population is obtained by applying the standard population strata-specific rates,  $p_1, \ldots, p_k$ , to the study population's as

$$E = \sum_{i=1}^{k} n_i p_i$$

The indirect adjusted rate is then

$$R_{\rm indirect} = C \frac{O}{E}$$

and O/E is the study population's standardized mortality ratio (SMR) if death is the event of interest or the standardized incidence ratio (SIR) for studies of disease incidence.

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The exact confidence interval is calculated for each estimated SMR by assuming a Poisson process as described in Breslow and Day (1980, 69–71). These are obtained by first calculating the upper and lower bounds for the confidence interval of the Poisson distributed observed events, O, say L and U respectively, and then computing:  $SMR_L = L/E$  and  $SMR_U = U/E$ .

#### Example

This example is borrowed from Kahn and Sempos (1989, 95–105). We want to compare 1970 mortality rates in California and Maine adjusting for age. Although we have age-specific population counts for the two states, we lack age-specific death rates. In this situation, direct standardization is not feasible. Since we have the US population census data for the same year, we can produce indirectly standardized rates for the two states.

The United States census, the standard population in this example, was entered into Stata and saved in popkahn.dta.

. use pop	okahn
-----------	-------

. list age pop deaths rate

	age	pop	deaths	rate
1.	<15	57900000	103062	.00178
2.	15-24	35441000	45261	.00128
з.	25-34	24907000	39193	.00157
4.	35-44	23088000	72617	.00315
5.	45-54	23220000	169517	.0073
6.	55-64	18590000	308373	.01659
7.	65-74	12436000	445531	.03583
8.	75+	7630000	736758	.09656

Note that the standard population contains for each age stratum the total number of individuals (pop) and both the age-specific mortality rate (rate) and the number of deaths. It is not necessary that the standard population contain all three. If you only have the age-specific mortality rate you can use the  $rate(ratevar_p \ crudevar_p)$  or  $rate(ratevar_p \ \#)$  options, where  $crudevar_p$  refers to the variable containing the total population's crude death rate or # is the total population's crude death rate.

Now let's look at the states data (study population).

. use Kahn					
. li	st				
	state	age	pop	death	
1.	California	<15	5524000	166285	
2.	California	15-24	3558000	166285	
з.	California	25-34	2677000	166285	
4.	California	35-44	2359000	166285	
5.	California	45-54	2330000	166285	
6.	California	55-64	1704000	166285	
7.	California	65-74	1105000	166285	
8.	California	75+	696000	166285	
9.	Maine	<15	286000	11051	
10.	Maine	15-24	168000		
11.	Maine	25-34	110000		
12.	Maine	35-44	109000		
13.	Maine	45-54	110000		
14.	Maine	55-64	94000		
15.	Maine	65-74	69000		
16.	Maine	75+	46000		

Note that for each state, the number of individuals in each stratum (age group) is contained in the variable pop. The death variable is the total number of deaths observed in the state during the year. It must have the same value for all observations in the group, such as for California, or it could be missing in all but one observation per group, such as in the case of Maine.

In terms of matching these two datasets, it is important that the strata variables have the same name in both datasets and ideally the same levels. If a level is missing from either dataset, the level will not be included in the standardization.

With these data loaded in memory (use Kahn), we now execute the command. We will use the print option to obtain the standard population's summary table and since we have both the standard population's age specific count and deaths, we will specify the popvars( $casevar_p \ popvar_p$ ) option. Alternatively, we could specify the rate(rate 0.00945) option since we know that 0.00945 is the US crude death rate for 1970. . is tdize death pop age using popkahn,  $\mbox{by}(\mbox{state})$   $\mbox{pop}(\mbox{deaths pop})$  print

Standard	Population
----------	------------

Stratum	Rate
<15	0.00178
15-24	0.00128
25-34	0.00157
35-44	0.00315
45-54	0.00730
55-64 65-74	0.01659 0.03583
75+	0.03656

#### Standard population's crude rate: 0.00945

-> state= Calif	ornia				
	Indirect St	andardization			
	Standard				
	Population	Observed	Cases		
Stratum	Rate	Population	Expected		
<15	0.0018	5524000	9832.72		
15-24	0.0013	3558000	4543.85		
25-34	0.0016	2677000	4212.46		
35-44	0.0031	2359000	7419.59		
45-54	0.0073	2330000	17010.10		
55-64	0.0166	1704000	28266.14		
65-74	0.0358	1105000	39587.63		
75+	0.0966	696000	67206.23		
Totals:		19953000	178078.73		
			Observed (	Cases:	166285
			SMR (Obs,	'Exp):	0.93
	SM	IR exact 95% C	onf. Interval:	0.9293,	0.9383]
			Crude	Rate:	0.0083
			Adjusted	Rate:	0.0088
		95% C	onf. Interval:	0.0088,	0.0089]
 -> state= Maine					
	Indirect St	andardization			
	Standard				
	Population	Observed	Cases		
Stratum	Rate	Population	Expected		
<15	0.0018	286000	509.08		
15-24	0.0013	168000	214.55		
25-34	0.0016	110000	173.09		
35-44	0.0031	109000	342.83		
45-54	0.0073	110000	803.05		
55-64	0.0166	94000	1559.28		
65-74	0.0358	69000	2471.99		
	0.0966	46000	4441.79		
75+	0.0000	10000			

(Continued on next page)

		SMR e:		SMR ( Conf. Interva Cr	rude Rate: sted Rate:	1.05 1.0707] 0.0111 0.0099
Summary of S	tudy Popul	ations (Rate	es):			
	Cases					
			0	Confidence		
California						
Maine	11051	0.011140	0.009931	[0.009747,	0.010118]	
Summary of Study Populations (SMR):						
	Cases	Case	S	Ex	act	
state		Expected		Confidenc	e Interval	
California Maine	166285		3 0.934	=	), 0.938271] 5, 1.070687]	

#### Acknowledgment

We gratefully acknowledge the collaboration of Dr. José Maria Pacheco from the Departamento de Epidemiologia, Faculdade de Saúde Pública/USP, Sao Paulo, Brazil and Dr. John L. Moran from The Queen Elizabeth Hospital, Woodville, Australia.

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Breslow, N. E. and N. E. Day. 1980. *Statistical Methods in Cancer Research*, Vol II. New York: Oxford University Press. Kahn, H. A. and C. T. Sempos. 1989. *Statistical Methods in Epidemiology*. New York: Oxford University Press.

snp1	4 Atv	vo-sample multivariate nonparametric test

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mv2snp performs the two-sample multivariate nonparametric test given by Leach (1991). The syntax of mv2snp is

mv2snp varlist, by(groupvar) verbose

where varlist consists of at least 2 variable names.

#### Options

by(groupvar) is not optional; it specifies the variable identifying the groups.

verbose is optional. If included, the vector of test statistics (U), variance-covariance matrix (V) and inverse of matrix NV will be reported (see below).

#### Methods and formulas

Data are ranked as a combined sample (as is carried-out for the univariate Mann–Whitney test) on each variable separately. The test statistic  $U^*$  is given as

$$U^* = U'(NV)^{-1}U$$

 $U^*$  is asymptotically distributed as  $\chi_p^2$  where p is the number of variables. Significant values of  $U^*$  indicate a difference between the two groups.

U is the vector of test statistics consisting of

$$U_i = \frac{R_i}{(N+1)} - \frac{n}{2}$$

for each variable, where  $R_i$  is the sum of ranks of the smaller group on the *i*th variable, *n* is the size of the smaller group, and *N* is the size of the combined sample. U' is the transpose of *U*. The variance-covariance matrix *V* is defined as having diagonal entries given by

$$v_{ii} = \frac{mn}{12N(N+1)}$$

and off-diagonal entries given by

$$v_{ij} = \frac{mn}{N^2(N-1)(N+1)^2} \left( \sum_{t=1}^N R_{it}R_{jt} - \frac{N(N+1)^2}{4} \right)$$

where m and n are the sizes of the two groups, N is the size of the combined sample (m + n) and  $\sum R_{it}R_{jt}$  is the sum of the cross-products of each pair of rankings across the combined sample. The matrix  $(NV)^{-1}$  is the inverse of NV where N is the combined sample size.

#### Example

The following example is that given by Leach (1991). Two psychological questionnaires (GHQ and IES) were given to two groups of policemen following a disaster at a soccer stadium. The first group (H) of 12 police were involved with the disaster; the second group (S) of 11 police were not involved.

The data are entered into Stata as three variables, ghq, ies and group:

. list ghq ies group

	$_{ghq}$	ies	group
1.	32	38	h
2.	34	17	h
з.	49	49	h
4.	33	45	h
5.	42	41	h
6.	49	49	h
7.	17	29	h
8.	48	43	h
9.	37	51	h
10.	48	48	h
11.	37	32	h
12.	22	51	h
13.	13	6	S
14.	13	14	S
15.	11	13	S
16.	21	36	S
17.	8	0	S
18.	24	23	S
19.	28	19	S
20.	9	0	S
21.	24	34	S
22.	22	7	S
23.	11	5	S

Running the test with the verbose option gives

```
. mv2snp ghq ies, by(group) verbose
Multi-variate non-parametric test (Leach, 1991)
Vector of test statistics, U
 -2.39583
 -2.41667
Variance-Covariance matrix, V
 0.01993 0.01561
 0.01561 0.01993
Inverse of matrix NV
  5.64142 -4.41781
 -4.41781 5.64142
 _____
                       _____
Test on variables: ghq ies
                      - GHQ Results
               ghq
                      - IES Results
               ies
Grouping variable: group
                      - Grouping Variable
Size of smaller group = 11
                           Number of variables = 2
Size of larger group = 12
                           U* =
                                  14.17
Combined sample size = 23
                           P = 0.00084
                    ____
```

Omitting the verbose option excludes output of vector U and matrices V and  $(NV)^{-1}$ .

## Saved results

mv2snp saves the following results:

S\_1 U\*
S\_2 Combined sample size
S\_3 Size of smaller sample
A Number of surjetles

S\_4 Number of variables

## Remarks

The grouping variable may be alpha or numeric and need not be named group. The data do not need to be sorted before applying mv2snp. When ranking the data, mv2snp does not exclude data values of zero. mv2snp does, however, perform casewise deletion for subjects with missing data because the nature of a multivariate test examines the correlation between variables; thus including data against which no correlation can be made will lead to erroneous results.

#### Reference

Leach, C. 1991. Nonparametric methods for complex data sets. In New Developments in Statistics for Psychology and the Social Sciences, Vol. 2. ed. P. Lovie and A. D. Lovie. British Psychological Society and Routledge.

#### STB categories and insert codes

Inserts in the STB are presently categorized as follows:

Gener	ral Categories:		
an	announcements	ip	instruction on programming
сс	communications & letters	os	operating system, hardware, &
dm	data management		interprogram communication
dt	datasets	qs	questions and suggestions
gr	graphics	tt	teaching
gr in	instruction	ZZ	not elsewhere classified
Statis	tical Categories:		
sbe	biostatistics & epidemiology	ssa	survival analysis
sed	exploratory data analysis	ssi	simulation & random numbers
sg	general statistics	SSS	social science & psychometrics
smv	multivariate analysis	sts	time-series, econometrics
snp	nonparametric methods	svy	survey sampling
sqc	quality control	sxd	experimental design
sqv	analysis of qualitative variables	SZZ	not elsewhere classified
srd	robust methods & statistical diagnostics		

In addition, we have granted one other prefix, stata, to the manufacturers of Stata for their exclusive use.

#### Guidelines for authors

The Stata Technical Bulletin (STB) is a journal that is intended to provide a forum for Stata users of all disciplines and levels of sophistication. The STB contains articles written by StataCorp, Stata users, and others.

Articles include new Stata commands (ado-files), programming tutorials, illustrations of data analysis techniques, discussions on teaching statistics, debates on appropriate statistical techniques, reports on other programs, and interesting datasets, announcements, questions, and suggestions.

A submission to the STB consists of

- 1. An insert (article) describing the purpose of the submission. The STB is produced using plain T<sub>F</sub>X so submissions using TEX (or LATEX) are the easiest for the editor to handle, but any word processor is appropriate. If you are not using TEX and your insert contains a significant amount of mathematics, please FAX (409-845-3144) a copy of the insert so we can see the intended appearance of the text.
- 2. Any ado-files, .exe files, or other software that accompanies the submission.
- 3. A help file for each ado-file included in the submission. See any recent STB diskette for the structure a help file. If you have questions, fill in as much of the information as possible and we will take care of the details.
- 4. A do-file that replicates the examples in your text. Also include the datasets used in the example. This allows us to verify that the software works as described and allows users to replicate the examples as a way of learning how to use the software.
- 5. Files containing the graphs to be included in the insert. If you have used STAGE to edit the graphs in your submission, be sure to include the .gph files. Do not add titles (e.g., "Figure 1: ...") to your graphs as we will have to strip them off.

The easiest way to submit an insert to the STB is to first create a single "archive file" (either a .zip file or a compressed .tar file) containing all of the files associated with the submission, and then email it to the editor at stb@stata.com either by first using uuencode if you are working on a Unix platform or by attaching it to an email message if your mailer allows the sending of attachments. In Unix, for example, to email the current directory and all of its subdirectories:

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