

# METAPERM: A SAS® Macro for Permutation Tests of Linear Models in Meta-analysis

Jeffrey D. Kromrey, Kristine Y. Hogarty, University of South Florida

Jeffrey D. Kromrey, University of South Florida, EDU 162, Tampa, FL 33620

## ABSTRACT

With the growing popularity of meta-analytic techniques to analyze and synthesize results across sets of empirical studies, has come concern about the sensitivity of traditional tests in meta-analysis to violations of assumptions. Such violations are particularly distressing because the tenability of such assumptions in primary studies is often impossible to evaluate. Permutation tests for linear models may provide a robust alternative to more traditional tests. A variety of permutation strategies for linear models have been proposed in the literature, and recent research (Kromrey & Hogarty, 2002; Hogarty & Kromrey, 2003) suggests that such approaches provide superior Type I error control when assumptions of normality and homogeneity of variance are violated. This paper presents a SAS macro that calculates four permutation tests of regression weights in linear models applied to meta-analysis (tests developed by Freedman & Lane, 1983; Ter Braak, 1992; Kennedy, 1995; and Manly, 1997), as well as the traditional parametric WLS test of these weights (Hedges & Olkin, 1985). The macro constructs a weighted linear model using a set of observed effect sizes, sample sizes, and values of hypothesized moderator variables that are supplied as inputs, and outputs the parameter estimates and probabilities obtained from the five testing procedures. The paper provides a demonstration of the SAS/IML code, sample output, and examples of applications in simulation studies.

## INTRODUCTION

Meta-analysis is a popular technique in many fields for statistically analyzing and synthesizing results across sets of empirical studies. Meta-analytic techniques provide a variety of models and procedures for pooling effect sizes across studies and for evaluating the effects of potentially moderating variables (Cooper & Hedges, 1994). However, both substantive concerns and statistical concerns about meta-analysis have been raised in the literature. For example, Fern and Monroe (1996) contended that the "interpretation and comparison of effect size across research studies is complicated by differences in substantive problems, theoretical perspectives, research methods, and researchers' goals" (p. 95). Further, concerns have been raised about the Type I error control and statistical power of meta-analytic tests when key assumptions are violated (Chang, 1993; Harwell, 1997; Hogarty & Kromrey, 1999). The sensitivity of traditional tests in meta-analysis to violations of assumptions is particularly distressing because the tenability of such assumptions (e.g., population normality, variance homogeneity) in the primary studies is often impossible to evaluate unless sufficient details are presented in reports

of primary studies (and such details are frequently not provided; Keselman, et al., 1998). The severity of the concerns that have recently been expressed in the literature suggests that alternative statistical approaches to meta-analysis are needed. Permutation tests may provide such a robust alternative.

## STATISTICAL TESTS FOR META-ANALYSIS

A fundamental purpose of meta-analysis is to differentiate between (a) collections of effect sizes that represent samples from a common population (i.e., having a common population effect size and differing from each other only because of sampling error) and (b) collections of effect sizes from different populations (i.e., having different population effect sizes). For the former situation, effects sizes may reasonably be pooled to provide both an estimate of the common population effect size and a confidence band around the estimate. For the latter situation, population effect sizes and confidence bands are estimated for each of the distinct populations.

*Traditional Parametric Tests.* Most meta-analysts use the family of Q tests (Hedges & Olkin, 1985) as tools for differentiating between these situations. According to Harwell (1997), from 1988 to 1995, of the 52 quantitative meta-analyses published by *Psychological Bulletin*, 60% employed Hedges' homogeneity test. The Q test of homogeneity evaluates the observed variability in sample effect sizes relative to the expected variability if all studies were sampled from a common population. Rejecting the null hypothesis of this Q test suggests that some (unspecified) moderator variable is present.

The logic of the Q test of homogeneity extends to the evaluation of between-group differences in mean effect sizes, a strategy that generalizes to the use of linear models for analysis of effect sizes (see, for example, Hedges, 1994; Raudenbush, 1994). That is, a linear model is fit to observed effect sizes:

$$d = \beta_0 + \beta_1 X_1 + \beta_2 X_2 + \dots + \beta_p X_p + e$$

where the Xs represent potential moderator variables, and the  $\beta_i$  represent the partial regression weights that relate the potential moderators to the observed effect sizes.

The parameters of this model are typically estimated using weighted least squares (WLS) or maximum likelihood (ML) methods that take into account differences in the sampling variability of the observed effect sizes. That is, the effect sizes are weighted by the inverse of the estimated sampling error.

As Harwell (1997) notes, despite the wide application of the Q test, the meta-analytic methodological literature provides little guidance in assessing the credibility of Q-test results if its assumptions are not tenable. Similar

concerns have been expressed by other researchers regarding the behavior of the Q-test when the underlying assumptions have been violated (Chandrashedaran & Walker, 1993; Chang, 1993; Wolf, 1990).

**Permutation Tests.** Permutation tests provide a promising approach to testing hypotheses in a variety of data structures. According to Good (1994), permutation tests are among the most powerful of statistical procedures available, offering robust alternatives in the face of violations of the assumptions of traditional parametric tests. The permutation strategy involves a comparison of the observed test statistic (e.g., differences in class mean effect sizes or estimated regression weights) with the set of values obtained through a rearrangement of the data. These rearrangements are repeated until a distribution is obtained for all possible permutations (an exact permutation test) or for a large, random sample of permutations (an approximate permutation test). This distribution of test statistics obtained from the permutations of the observed data provides an empirical sampling distribution with which to compare the observed test statistic. The permutation strategy holds promise for providing a method of testing hypotheses in meta-analysis, avoiding the problems of poor Type I error control and power associated with the Q test.

The application of permutation tests to partial regression weights is more challenging than the application to bivariate relationships. In bivariate models such as a zero-order correlation or bivariate regression, any pairing of the observed  $x$  and  $y$  values is equally probable under the null hypothesis. Thus, the elements of the  $y$  vector (or equivalently, the  $x$  vector) may be directly permuted to construct a statistically valid test of the null hypothesis. With multiple regression applications, and the construction of permutation tests for partial regression weights, such naïve permutation is not valid because the observed  $y$  values are a function of the set of regressors, rather than a single regressor (i.e., the observed  $y$  values are not exchangeable under the null hypothesis that a particular partial regression weight is zero). The permutation needs to be conducted on that part of  $y$  and that part of  $x_i$  that are unrelated to the other regressors. Thus the focus is on partial correlation coefficients to derive a test for the regression weights. Differences among permutation methods suggested in the literature reflect differences in how these partial correlations should be obtained when conducting a permutation test.

Consider the typical squared partial correlation coefficient (in this example, the correlation between  $y$  and  $z$ , while controlling for  $x$ ) as the correlation between two residuals:

$$r_{yz.x}^2 = \frac{\sum (res_{y.x} res_{z.x})^2}{\sum res_{y.x}^2 \sum res_{z.x}^2}$$

where  $res_{y.x}$  is the residual of  $y$  after removing  $x$  (note that  $x$  may be a single variable or a set of regressors). This squared partial correlation is used as the test statistic for testing the partial regression weight of  $z$  in the equation with the  $x$  variables. The differences among the four methods reflect differences in the statistics used to construct the permutation distribution.

The four methods will be illustrated using a simple example of a criterion variable ( $y$ ) and a set of regressors

[ $x$ ,  $z$ ]. The regressors have been partitioned into a set  $x$  and a single regressor  $z$ . In meta-analytic applications,  $y$  represents the observed effect sizes and the regressors are potential moderating variables. For a given sample of observed values of  $y$ ,  $z$  and  $x$ ,  $y$  is regressed on  $x$  to obtain the residuals  $res_{y.x}$  and the predicted values  $y'$ . Subsequently,  $z$  is regressed on  $x$  to obtain the residuals  $res_{z.x}$ .

The permutation distribution suggested by Freedman and Lane (1983) is constructed by permuting the residuals  $res_{y.x}$  and adding them to the predicted values  $y'$  to construct a new set of  $y$  variables (these new variables are represented as  $y^p$  because they are not actual data that were observed, but are constructed from a single permutation of the observed data). Now, these  $y^p$  values may be regressed on  $x$  to obtain another set of residuals that are unique to this permutation of the data ( $res_{FL_{y,x}}$ ) and the squared partial correlation coefficient for this permutation is obtained as:

$$FL_{yz.x}^2 = \frac{\sum (res_{FL_{y,x}} res_{z,x})^2}{\sum (res_{FL_{y,x}})^2 \sum res_{z,x}^2}$$

Note that the residuals involving  $z$  and  $x$  have not changed in the permutation – their values are constant across the set of permutations.

The permutation distribution suggested by Kennedy (1995) is also constructed by permuting the residuals  $res_{y,x}$ , but they are not recombined with the original predicted values. Rather, these permuted residuals are entered directly in the calculation of the squared partial correlation:

$$KEN_{yz.x}^2 = \frac{\sum (res_{y,x} res_{z,x})^2}{\sum res_{y,x}^2 \sum res_{z,x}^2}$$

The only value that will change across permutations is the numerator of this formula, because each permutation will result in new pairings of the two residuals, while the sum of the squared residuals remains constant.

Manly (1997) suggested that the original observed  $y$  values may be permuted, and the regression of these permuted  $y$  values on  $x$  may be obtained, providing residuals ( $res_{MN_{y,x}}$ ). These residuals, which will be unique for each permutation of the  $y$  vector, are used to compute the partial correlation:

$$MN_{yz.x}^2 = \frac{\sum (res_{MN_{y,x}} res_{z,x})^2}{\sum (res_{MN_{y,x}})^2 \sum res_{z,x}^2}$$

Finally, Ter Braak (1992) suggested a permutation distribution that is similar to the Freedman and Lane approach, except that the residuals being permuted are obtained from regressing  $y$  on both  $z$  and  $x$  simultaneously (called the full model residuals). For a given sample of observed values of  $y$ ,  $z$  and  $x$ , the observed values of  $y$  are regressed on  $x$  and  $z$  simultaneously to obtain the residuals  $res_{y,xz}$ . The permutation distribution suggested by Ter Braak is constructed by permuting these residuals, then regressing them on  $x$  (only) to obtain another set of residuals that are unique to this permutation of the data ( $res_{TB_{y,x}}$ ). The squared partial correlation coefficient for

this permutation is obtained as

$$TB_{-r_{yz,x}^2} = \frac{\sum (res_{-TB_{y,x}} res_{z,x})^2}{\sum (res_{-TB_{y,x}})^2 \sum res_{z,x}^2}$$

In all four methods, the observed value of the squared partial correlation is used as the test statistic, however, the four methods yield different permutation distributions against which the value is evaluated to obtain probability statements. Previous investigations of these approaches to testing weights in linear models (e.g., Anderson & Legendre, 1999; Anderson & Robinson, 2001) suggest that they yield nearly identical asymptotic distributions, but evidence substantial differences in finite samples. Further, these approaches have not been investigated in the context of weighted estimation in linear models such as that presented by meta-analysis.

## MACRO METAPERM

A SAS/IML macro was designed to compute the tests of regression weights using the permutation methods as well as the standard WLS approach. The macro was developed to provide researchers with an easily accessible tool for conducting robust tests in meta-analysis. Inputs to the macro include the name of the SAS dataset containing the effect sizes and information about study characteristics, variable names in the SAS dataset for the effect sizes and sample sizes in the primary studies, the number of regressor variables to be included in the model, the number of permutations to execute and the alpha level for the tests of weights. The names of the regressor variables in the SAS dataset must be X1, X2, X3, etc.

```

*-----+
Inputs to the macro:
dsn: name of SAS data set with effect sizes and study
characteristics
eff_size: name of variable that represents the observed effect
sizes
n1, n2: name of variable that represents sample sizes in the
two groups
n_x: number of regressor variables for the model
n_perms: number of permutations to generate
alpha: nominal alpha level for tests
+-----+

%macro METAPERM (dsn,eff_size,n_1,n_2,n_x, n_perms,alpha);
proc iml;
*-----+
Subroutine to randomly permute the rows in a matrix
Input: ORIG_MTX
Output: PERM_MTX
+-----+
start permute(orig_mtx,perm_mtx);
L = nrow(orig_mtx);
W = ncol(orig_mtx);
perm_mtx=J(L,W,0);

Ranvec=J(L,1,0);
do i = 1 to L;
Ranvec[i,1] = ranuni(0);
end;
Rankvec = rank(ranvec);

do i = 1 to L;

```

```

perm_mtx[i,] = orig_mtx[rankvec[i,1],];
end;
Finish;

```

```

*-----+
Subroutine to calculate the WLS tests of fixed-effects
models and residuals to use in permutations.
Inputs to the subroutine are
di_vec - column vector of effect sizes (d)
n_vec - matrix (k X 2) of sample sizes corresponding to each
effect size
X_Matrix - Matrix of potential moderator variables

Outputs are
Vi - reciprocals of variances, used for weights
B_wls - regression weights for full model
SE_B - Standard errors of the regression weights
Resid_wls - Matrix of weighted least squares residuals, first
column from full model, other columns
from leaving out each regressor
Resid_ols - Matrix of ordinary least squares residuals, each
regressor predicted by others
Partcorr - vector of partial correlations of Y with each
regressor
Pred_wls - Matrix of wls predicted values
Pred_ols - Matrix of ols predicted values of each regressor
+-----+
start calcreq(di_vec,n_vec,X_Matrix,Vi,B_wls,SE_B,
Resid_wls,Resid_ols,partcorr,pred_wls,pred_ols);

* calculate variance for each effect size;
k = nrow(di_vec);
var_di=J(k,1,0);
Vi=J(k,1,0);
do i = 1 to k;
var_di[i,1] = ((n_vec[i,1]+n_vec[i,2])/(n_vec[i,1]*n_vec[i,2]))
+
((di_vec[i,1]**2)/(2*(n_vec[i,1]+n_vec[i,2])));
Vi[i,1]=var_di[i,1]**-1;
end;
*-----+
Weighted least squares estimation using Vi as weights
+-----+
* All regressors;
X = X_Matrix;
B_wls = INV(X**DIAG(Vi)*X)*X**DIAG(Vi)*di_vec;
Resid_wls = di_vec - X*B_wls;
Pred_wls = X*B_wls;
cov_b = INV(X**DIAG(Vi)*X);
SE_B = SQRT(vecdiag(cov_b));

* Leaving out each regressor;
N_X = ncol(X);
do col = 2 to N_X;
do col2 = 1 to N_X;
if col2 = 1 then temp_X = X[,1];
if col2 > 1 then do;
if col2 ^= col then temp_X = temp_X||X[,col2];
end;
end;

Target_X = X[,col];
B_wls_T = INV(temp_X**DIAG(Vi)*temp_X)*
temp_X**DIAG(Vi)*di_vec;
Resid_wls = Resid_wls || (di_vec - temp_X*B_wls_T);
Pred_wls = Pred_wls || (temp_X*B_wls_T);
B_ols_T = INV(temp_X**temp_X)*temp_X**Target_X;
if col = 2 then do;
Resid_ols = (Target_X - temp_X*B_ols_T);
Pred_ols = temp_X*B_ols_T;
end;
if col > 2 then do;
Resid_ols = Resid_ols || (Target_X - temp_X*B_ols_T);
Pred_ols = pred_ols || (temp_X*B_ols_T);

```

```

end;
end;

* Compute partial correlations;
partcorr = J(N_X-1,1,0);
do col = 1 to N_X - 1;
  num = 0;
  denom1 = 0;
  denom2 = 0;
  do row = 1 to k;
    num = num + Vi[row,1]#(resid_wls[row,col+1] #
      resid_ols[row,col])##2;
    denom1 = denom1 + Vi[row,1]# resid_wls[row,col+1]##2;
    denom2 = denom2 + resid_ols[row,col]##2;
  end;
  partcorr[col,1] = num/(denom1 # denom2);
end;
finish;

```

### Main program

```

use &dsn;
read all var{&eff_size} into di_vec;
read all var{&n_1} into n1;
read all var{&n_2} into n2;
n_vec = n1||n2;
free n1 n2;
%do i = 1 %to &n_x;
  read all var{x&i} into temp_x;
  if &i = 1 then do;
    X_Matrix = temp_x;
  End;
  If &i > 1 then do;
    X_matrix = X_matrix || temp_x;
  End;
%end;
k = nrow(X_matrix);
X_matrix = J(k,1,1)||X_Matrix;

run calcreq(di_vec,n_vec,X_Matrix,Vi,B_wls,SE_B,
  Resid_wls,Resid_ols,partcorr,pred_wls,pred_ols);

```

### Normal theory WLS tests of regression weights

```

Xs = ncol(X_Matrix) - 1;
PROB_FEZ = J(Xs,1,0);
do ii = 2 to Xs+1;
  FE_Z=B_wls[ii,1]/SE_B[ii,1];
  PROB_FEZ[ii-1,1] = 2#(1-probnorm(abs(FE_Z)));
end;
Resid_wls = resid_wls||di_vec;

```

```

FL = J(Xs,1,0);
Ken = J(Xs,1,0);
MN = J(Xs,1,0);
TB = J(Xs,1,0);

```

```

do perm = 1 to &n_perms;

  run permute(Resid_wls,Perm_Res);

```

### Freedman and Lane Method

```

do XX = 2 to Xs+1;
  new_y = Perm_Res[XX] + pred_wls[XX];
  run calcreq(new_y,n_vec,X_Matrix,j7,j1,j2,j3,j4, pcorr_i,j5,j6);
  if pcorr_i[XX-1,1]>partcorr[XX-1,1] then FL[XX-1,1] = FL[XX-
    1,1] + 1;
  free new_y j1 j2 j3 j4 j5 j6 j7;
end;

```

### Kennedy Method

```

+-----+;
K_corr = J(Xs,1,0);
do col = 1 to Xs;
  num = 0; denom1 = 0; denom2 = 0;
  do row = 1 to k;
    num = num + Vi[row,1]#(Perm_Res[row,col+1] #
      resid_ols[row,col])##2;
    denom1 = denom1 + Vi[row,1]#Perm_Res[row,col+1]##2;
    denom2 = denom2 + resid_ols[row,col]##2;
  end;
  K_corr[col,1] = num/(denom1 # denom2);
  if K_corr[col,1]>partcorr[col,1] then Ken[col,1] =
    Ken[col,1] + 1;
end;

```

### Manly Method

```

+-----+;
new_y = Perm_Res[5];
run calcreq(new_y,n_vec,X_Matrix,j7,j1,j2,j3,j4,pcorr_i,j5,j6);
free new_y j1 j2 j3 j4 j5 j6 j7;
do XX = 2 to Xs+1;
  if pcorr_i[XX-1,1]>partcorr[XX-1,1] then MN[XX-1,1] = MN[XX-
    1,1] + 1;
end;

```

### Ter Braak Method

```

+-----+;
new_y = Perm_Res[1] + pred_wls[1];
run calcreq(new_y,n_vec,X_Matrix,j7,j1,j2,j3,j4,pcorr_i,j5,j6);
free new_y j1 j2 j3 j4 j5 j6 j7;
do XX = 2 to Xs+1;
  if pcorr_i[XX-1,1]>partcorr[XX-1,1] then TB[XX-1,1] = TB[XX-
    1,1] + 1;
end;
end; * end the permutation loop;

```

```

do z = 1 to Xs;
  FL[z,1] = FL[z,1]/&n_perms;
  Ken[z,1] = Ken[z,1]/&n_perms;
  MN[z,1] = MN[z,1]/&n_perms;
  TB[z,1] = TB[z,1]/&n_perms;
end;
nperms = &n_perms;

```

### Create Output Table

```

+-----+;
file print;
put @1 'Linear Model for Meta Analysis' /
@1 'Tests of Regression Weights' /
@1 '-----' /
@1 'N of Studies:' @48 k 8. /
@1 'Number of Permutations:' @48 nperms 8. //
@1 '-----' /
@1 ' Probabilities Under Ho: beta = 0' /
@1 '-----' /
@1 ' Parameter Freedman Ter/
@1 'Regressor Estimate WLS Lane Kennedy Manly
Braak' /
@1 '-----';
do i = 1 to &n_x;
  p_FL = FL[i,1];
  if p_FL < &alpha then flagFL = '*'; else flagFL = ' ';
  p_KN = Ken[i,1];
  if p_KN < &alpha then flagKN = '*'; else flagKN = ' ';
  p_MN = MN[i,1];
  if p_MN < &alpha then flagMN = '*'; else flagMN = ' ';
  p_TB = TB[i,1];
  if p_TB < &alpha then flagTB = '*'; else flagTB = ' ';
  p_WS = PROB_FEZ[i,1];
  if p_WS < &alpha then flagWS = '*'; else flagWS = ' ';
  beta = B_wls[i+1,1];

```



greatest power with few studies in the meta-analysis ( $k = 10$ ). The Ter Braak test was conservative across most of the conditions examined, resulting in very low power estimates. However, for conditions in which the normal theory assumptions hold, the WLS test was notably more powerful than any of the permutation tests.

## CONCLUSIONS

Meta-analysis has become increasingly important for the synthesis of research results in a variety of fields, including education, the behavioral sciences and medicine. The accuracy of inferences derived from meta-analysis depends upon the appropriate application of statistical tools. As the use of meta-analytic methods becomes more commonplace, researchers must remain mindful of the limitations of certain estimates. Permutation tests can provide a robust alternative to traditional parametric tests for meta-analysis when critical assumptions are violated.

The macro METAPERM is provided to facilitate researchers' calculation and use of four permutation tests for testing moderating effects in meta-analysis. Although the macro, as provided, is limited to the analysis of standardized mean differences as effect sizes (i.e., Cohen's  $d$ ), the code is easily modified for the analysis of other indices of effect magnitude. For example, the analysis of Pearson Product Moment Correlation Coefficients requires a modification of the calculation of the variance in these effect sizes, and the incorporation of Fisher's  $z$  transformation to normalize the sampling distribution of  $r$ .

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## CONTACT INFORMATION

Your comments and questions are valued and encouraged. Please contact Jeff Kromrey at:  
University of South Florida  
4202 East Fowler Ave. EDU 162  
Tampa, FL 33620  
Work Phone: 813-974-5739  
Fax: 813-974-4495  
Email: kromrey@tempest.coedu.usf.edu