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The Stata Journal is published quarterly by the Stata Press, College Station, Texas, USA.

Address changes should be sent to the *Stata Journal*, StataCorp, 4905 Lakeway Drive, College Station, TX 77845, USA, or emailed to sj@stata.com.



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# A short guide and a forest plot command (ipdforest) for one-stage meta-analysis

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**Abstract.** In this article, we describe a new individual patient data meta-analysis postestimation command, ipdforest. The command produces a forest plot following a one-stage meta-analysis with xtmixed or xtmelogit. (These commands have been renamed in Stata 13 to mixed and meqrlogit, respectively; ipdforest is currently not compatible with the new names.) The overall effect is obtained from the preceding mixed-effects regression and the study effects from linear or logistic regressions on each study, which are executed within ipdforest. Individual patient data meta-analysis models with Stata are discussed.

 ${\sf Keywords:}$ st<br/>0309, ipdforest, meta-analysis, forest plot, individual patient data, IPD, one-stage

# 1 Introduction

Meta-analysis, the methodology that allows results from independent studies to be combined, is usually a two-stage process. First, the relevant summary effect statistics are extracted from published articles on the included studies. These are then combined into an overall effect estimate using a suitable meta-analysis model (Harris et al. 2008; Kontopantelis and Reeves 2010). However, problems often arise when an article does not report all the statistical information required as input for the meta-analysis (for example, it fails to provide a variance estimate for the outcome measure); reports a statistic other than the effect size (such as a t-value or p-value) that needs to be transformed with a loss of precision; or provides a sample too clinically heterogeneous for the study to be included in the meta-analysis (Kontopantelis and Reeves 2009).

When individual patient data (IPD) from each study are available, meta-analysts can avoid these problems when estimating study effects; outcomes can be easily standardized, while clinical heterogeneity can be addressed, at least partially, with subgroup analyses and patient-level covariate control. Furthermore, when IPD data are available, meta-analysts can use a mixed-effects regression model to combine information across studies in a single stage. This is recognized as the best approach for performing an IPD meta-analysis, with the two-stage method being at best equivalent in certain scenarios (Mathew and Nordström 2010). Despite these advantages of the one-stage approach, one obvious advantage of twostage meta-analysis is the ability to convey information graphically through a forest plot. Because study effects have been calculated or extracted in the first stage of the process, they and their respective confidence intervals can be used to demonstrate the relative strength of the intervention in each study and across all the studies. Forest plots are informative, easy to follow, and particularly useful for readers with little or no experience in meta-analysis methods. It is not surprising, then, that they have become a key feature of meta-analysis and are always presented when two-stage meta-analyses are performed.

However, under a one-stage meta-analysis model, only the overall effect is calculated, not individual study effects; thus creating a forest plot is not straightforward. A search by the authors failed to identify one-stage meta-analysis forest-plot modules in any general or meta-analysis specialist statistical package. We attempt to address this gap in Stata with the **ipdforest** command.

This article is divided into two sections. In the first section, we describe IPD metaanalysis models and their implementation in Stata with available mixed-effects models. In the second section, we describe the **ipdforest** command in detail and provide an example.

# 2 Individual patient data meta-analysis

A description of IPD meta-analysis methods for continuous and binary outcomes has been provided by Higgins et al. (2003) and Turner et al. (2000), respectively. Although we will only explore a representative selection of linear random-effects models in Stata (using the xtmixed command), application to the logistic case using xtmelogit should be straightforward. Let us assume IPD from a group of studies. For each trial, we have the exposure variable, which is continuous or binary (for example, control or intervention group membership), and baseline and follow-up data for the continuous outcome and covariates. We will also assume that both the outcome measure and any covariates have been measured in the same way across studies and that, therefore, standardization is not required. In the models that follow, in general, we denote a fixed effect by  $\gamma$  and a random effect by  $\beta$ .

Possibly the simplest approach is to assume that there is a common intercept across studies and that baseline is a fixed effect but to allow the treatment effect to vary at random across studies. Thus we have

$$Y_{ij} = \gamma_0 + \beta_{1j} \operatorname{group}_{ij} + \gamma_2 Y_{ij} + \epsilon_{ij}$$
  
$$\beta_{1j} = \gamma_1 + u_{1j}$$
(1a)

and

$$\epsilon_{ij} \sim N(0, \sigma_j^2)$$

$$u_{1i} \sim N(0, \tau_1^{-2})$$
(1b)

where *i* is the patient; *j* is the study;  $Y_{ij}$  is the outcome for patient *i* in study *j*;  $\gamma_0$  is the fixed common intercept;  $\beta_{1j}$  is the random treatment effect for study *j*;  $\gamma_1$  is the mean treatment effect; group<sub>*ij*</sub> is the exposure for patient *i* in study *j*;  $\gamma_2$  is the fixed baseline effect;  $Y_{ij}$  is the baseline score for patient *i* in study *j*;  $u_{1j}$  is the random treatment effect for study *j* (shifting the regression line up or down by study);  $\tau_1^2$  is the between-study variance;  $\epsilon_{ij}$  is the error term for patient *i* in study *j*; and  $\sigma_j^2$  is the within-study variance for study *j*.

However, the common intercept and fixed baseline assumptions are difficult to justify, and such a model should be approached with caution—if at all. A more accepted model allows for different fixed intercepts and fixed baseline effects for each study:

$$\dot{Y}_{ij} = \gamma_{0j} + \beta_{1j} \operatorname{group}_{ij} + \gamma_{2j} Y_{ij} + \epsilon_{ij} 
\beta_{1j} = \gamma_1 + u_{1j}$$
(2)

where  $\gamma_{0j}$  is the fixed intercept for study j and  $\gamma_{2j}$  is the fixed baseline effect for study j.

Another possibility, although contentious (Whitehead 2002), is to assume that study intercepts are random, as in a multicenter study; for example,

$$\dot{Y}_{ij} = \beta_{0j} + \beta_{1j} \operatorname{group}_{ij} + \gamma_{2j} Y_{ij} + \epsilon_{ij} 
\beta_{0j} = \gamma_0 + u_{0j} 
\beta_{1j} = \gamma_1 + u_{1j}$$
(3a)

In this case, it is probably wiser to assume a nonzero correlation  $\rho$  between the random effects:

$$\epsilon_{ij} \sim N(0, \sigma_j^2)$$

$$u_{0j} \sim N(0, \tau_0^2)$$

$$u_{1j} \sim N(0, \tau_1^2)$$

$$\operatorname{cov}(u_{0j}, u_{1j}) = \rho \tau_0 \tau_1$$
(3b)

The baseline could also have been modeled as a random effect, and we could have allowed for nonzero correlations between the three random effects, thus complicating (3) further:

$$\dot{Y}_{ij} = \beta_{0j} + \beta_{1j} \operatorname{group}_{ij} + \beta_{2j} Y_{ij} + \epsilon_{ij} 
\beta_{0j} = \gamma_0 + u_{0j} 
\beta_{1j} = \gamma_1 + u_{1j} 
\beta_{2j} = \gamma_2 + u_{2j}$$
(4a)

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with effects

$$\begin{aligned} \epsilon_{ij} &\sim N(0, \sigma_j^2) \\ u_{0j} &\sim N(0, \tau_0^2) \\ u_{1j} &\sim N(0, \tau_1^2) \\ u_{2j} &\sim N(0, \tau_2^2) \\ \cos(u_{0j}, u_{1j}) &= \rho_1 \tau_0 \tau_1 \\ \cos(u_{0j}, u_{2j}) &= \rho_2 \tau_0 \tau_2 \\ \cos(u_{1j}, u_{2j}) &= \rho_3 \tau_1 \tau_2 \end{aligned}$$
(4b)

In some cases, the focus might be on interactions. For example, if we assume a continuous and standardized variable X, we can expand (2) to include fixed effects, in this instance, for both X and its interaction with the treatment:

$$\dot{Y}_{ij} = \gamma_{0j} + \beta_{1j} \operatorname{group}_{ij} + \gamma_{2j} Y_{ij} + \gamma_3 X_{ij} + \gamma_4 \operatorname{group}_{ij} X_{ij} + \epsilon_{ij} 
\beta_{1j} = \gamma_1 + u_{1j}$$
(5)

If we consider Yfin and Ybas as representing the outcome and baseline, respectively, the exposure variable group, and the study identifier studyid for four studies, we can implement the models described above by using xtmixed.

Model (1): Fixed common intercept; random treatment effect; fixed effect for baseline.

. xtmixed Yfin i.group Ybas || studyid:group, nocons

The **nocons** option suppresses estimation of the intercept as a random effect.

Model (2): Fixed study-specific intercepts; random treatment effect; fixed study-specific effects for baseline (where Ybas'i'=Ybas if studyid='i' and equals 0 otherwise).

. xtmixed Yfin i.group i.studyid Ybas1 Ybas2 Ybas3 Ybas4 || studyid:group, > nocons

Model (3): Random study intercept; random treatment effect; fixed study-specific effects for baseline.

. xtmixed Yfin i.group Ybas1 Ybas2 Ybas3 Ybas4 || studyid:group, cov(uns)

Model (4): Random study intercept; random treatment effect; random effect for baseline.

. xtmixed Yfin i.group Ybas || studyid:group Ybas, cov(uns)

In general, a covariate (or an interaction term) can be modeled as a fixed or random effect, but in the latter case, the complexity of the model increases and nonconvergence issues are more likely to be encountered. If we also consider patient covariate **age** and its interaction with the treatment effect, then (5) will be

. xtmixed Yfin i.group i.studyid Ybas1 Ybas2 Ybas3 Ybas4 age i.group#c.age > || studyid: group, nocons

Or alternatively, age can be modeled as a random effect:

```
. xtmixed Yfin i.group i.studyid Ybas1 Ybas2 Ybas3 Ybas4 age i.group#c.age
> || studyid: group age, nocons
```

## 3 The ipdforest command

#### 3.1 Syntax

```
ipdforest varname [, re(varlist) fe(varlist) fets(namelist) ia(varname) auto
    label(varlist) or gsavedir(string) gsavename(string) eps gph
    export(string)]
```

where *varname* is the exposure variable, continuous or binary (for example, intervention or control).

## 3.2 Options

- re(varlist) specifies covariates to be included as random factors. For each covariate specified, a different regression coefficient is estimated for each study.
- fe(*varlist*) specifies covariates to be included as fixed factors. For each covariate specified, the respective coefficient in the study-specific regressions is fixed to the value returned by the multilevel regression.
- fets(namelist) specifies covariates to be included as study-specific fixed factors (that is, by using the estimated study fixed effects from the main regression in all individual study regressions). Only baseline scores and study identifiers can be included. For each covariate specified, the respective coefficient in the study-specific regressions is fixed to the value returned by the multilevel regression for the specific study. For study-specific intercepts, the study identifier (not in factor-variable format, for example, studyid) or the stub of the dummy variables (for example, studyid\_ when dummy study identifiers are studyid\_1 studyid\_3, etc.) would be included. For study-specific baseline scores, only the stub of the dummy variables is accepted (for example, dept0s\_ when dummy study baseline scores are dept0s\_1 dept0s\_3, etc.).

- ia(varname) specifies covariates for which the interaction with the exposure variable will be calculated and displayed. The covariate should also be specified as a fixed, random, or study-specific fixed effect. If binary, the command will provide two sets of results, one for each group. If categorical, it will provide as many sets of results as there are categories. If continuous, it will provide one set of results for the main effect and one for the interaction. Although the command will allow a variable to be interacted with the exposure variable as a fixed or study-specific fixed effect, the variable necessarily will be included as a random effect in the individual regressions (it will not run a regression with the interaction term only; the main effects must be included as well). Therefore, although the overall effect will differ between a model with a fixed-effects interacted variable and a random-effects one, the individual study effects will be identical across the two approaches.
- auto allows ipdforest to automatically detect the specification of the preceding model. This option cannot be issued along with options re(), fe(), fets(), or ia(). The auto option will work in most situations, but it comes with certain limitations. It uses the returned command string of the preceding command, which is effectively constrained to 244 characters; therefore, the **auto** option will return an error if ipdforest follows a very wide regression model—in such a situation, only the manual specification can be used. In addition, the variable names used in the preceding model must follow certain rules: 1) fixed-effects covariates (manually with option fe()) must not contain underscores; 2) for study-specific intercepts (manually with option fets()), factor-variable format is allowed or a *varlist* (for example, cons\_2cons\_16), but each variable must contain a single underscore followed by the study number (not necessarily sequential); and 3) for study-specific baseline scores (manually with option fets()), each variable must contain a single underscore followed by the study number (again, not necessarily sequential). There are no restrictions for random-effects covariates (manually with option re()). For interactions (manually with option ia(), the factor-variable notation should be preferred (for example, i.group#c.age) and, alternatively, the older xi: notation. Interactions expanded to dummy variables cannot be identified with the **auto** option, and only the manual specification should be used in this case. Variables whose names start with an \_I and contain a capital X will be assumed to be expanded interaction terms, and if detected in the last model, ipdforest will terminate with a syntax error.
- label(varlist) specifies labels for the studies. Up to two variables can be specified and converted to strings. If two variables are specified, they will be separated by a comma. Usually, the author names and the year of study are selected as labels. If label() is not specified, the command automatically uses the value labels of the numeric cluster variable, if any, to label the forest plot. Either way, the final string is truncated to 30 characters.
- or reports odds ratios instead of coefficients. It can only be used following the execution of xtmelogit.
- gsavedir(*string*) specifies the directory in which to save the graph, if different from the active directory.

gsavename(string) specifies the optional name prefix for the graph. Graphs are saved as gsavename\_graphname.gph or gsavename\_graphname.eps, where graphname includes a description of the summary effect (for example, main\_group for the main effect if group is the exposure variable).

eps saves the graph in .eps format instead of the default .gph.

- gph saves the graph in .gph format. gph is the default. Use it to save in both formats: including only the eps option will save the graph in .eps format only.
- export(string) exports the study identifiers, weights, effects, and standard errors in a Stata dataset (named after string). It is provided for users who wish to use other commands or software to draw the forest plots.

#### 3.3 Stored results

ipdforest stores the following in r():

Scalars

r(Isq) r(Hsq)	heterogeneity measure $I^2$ heterogeneity measure $H^2_M$	r(eff1pe_ov) r(eff1se_ov)	overall effect estimate standard error of the overall
r(tausq)	$\hat{\tau}^2$ , between-study variance	1(011100_01)	effect
	estimate	r(eff1pe_st <i>i</i> )	effect estimate for study $i$
r(tausqlo)	$\hat{\tau}^2$ , lower 95% confidence	r(eff1se_st <i>i</i> )	standard error of the effect
	interval		for study $i$
r(tausqup)	$\hat{\tau}^2$ , upper 95% confidence		
	interval		

If an interaction with a continuous variable is included in the model, it also stores the following:

Scalars

r(eff2pe_ov)	overall interaction effect	r(eff2pe_st <i>i</i> )	interaction effect estimate for
	estimate		study $i$
r(eff2se_ov)	standard error of the overall	r(eff2se_st <i>i</i> )	interaction effect standard
	interaction effect		error for study $i$

If the interaction variable is binary, the first set of results corresponds to the effects for the first category of the binary (for example, sex = 0) and the second set for the second category (for example, sex = 1). If the variable is categorical, the command returns as many sets of effect results as there are categories (with each set corresponding to one category). Estimation results from xtmixed or xtmelogit in e() are restored after the execution of ipdforest.

### 3.4 Methods

The ipdforest command is issued following a random-effects IPD meta-analysis conducted using a linear (xtmixed) or logistic (xtmelogit) two-level regression with patients nested within studies. The command provides a meta-analysis summary table and a forest plot. Study effects are calculated within ipdforest, while the overall effect and variance estimates are extracted from the preceding regression. The default estimation methods for xtmixed and xtmelogit are restricted maximum likelihood and maximum likelihood, respectively. A description of these methods is beyond the scope of this article.

ipdforest estimates individual study effects and their standard errors by using one-level linear or logistic regression analyses. Following xtmixed, regress is used, and following xtmelogit, logit is used for each study in the meta-analysis. The ipdforest command controls these regressions for fixed- or random-effects covariates that were specified in the preceding two-level regression. The user has full control over the covariates to be included in the ipdforest command, including their specification as fixed or random effects. However, we strongly recommend using the same specification as in the preceding xtmixed or xtmelogit command because the reported overall effect and its confidence interval is taken from that model.

In the estimation of individual study effects, ipdforest controls for a random-effects covariate (that is, allowing the regression coefficient to vary by study) by including the covariate as an independent variable in each regression. Control for a fixed-effects covariate (where the regression coefficient is assumed constant across studies and is given by the coefficient estimated under xtmixed or xtmelogit) is a little more complex. Because it is not possible to specify a fixed value for a regression coefficient under regress, the continuous outcome variable is adjusted by subtracting the contribution of the fixed covariates to its values prior to analysis. For a binary outcome, the equivalent is achieved using the offset option in logit. Patient weights are uniform; therefore, each study's weight is the ratio of its participants over the total number of participants across all studies.

Between-study variability in the treatment effect, known as heterogeneity, arises from differences in study design, quality, outcomes, or populations and needs to be accounted for in the meta-analysis model when present. Heterogeneity is usually reported in the form of measures or tests that compare the between- and within-study variance estimates. For continuous outcomes, **ipdforest** reports two heterogeneity measures,  $I^2$  and  $H_M^2$ , based on the **xtmixed** output.  $I^2$  values of 25%, 50%, and 75% correspond to low, moderate, and high heterogeneity, respectively (Higgins et al. 2003), while  $H_M^2$  takes values in the  $[0, +\infty)$  range with 0 indicating perfect homogeneity (Mittlböck and Heinzl 2006). We have not attempted to calculate an IPD version of Cochran's Q, the orthodox  $\chi^2_{k-1}$  homogeneity test, considering its poor performance when the number of studies k is small (Hardy and Thompson 1998). For binary outcomes, an estimate of the within-study variance is not reported under **xtmelogit**, and hence, heterogeneity measures cannot be computed. The between-study variance estimate  $\hat{\tau}^2$  and its confidence interval are reported under both models.

Fixed-effects meta-analysis models are widely used when heterogeneity is very low or 0. However, a more conservative approach is to take account of even low levels of between-study variability by adopting a random-effects model (Hunter and Schmidt 2000). When between-study variance is estimated to be close to 0, results with the two approaches converge. Therefore, although ipdforest is a postestimation command for random-effects IPD meta-analysis, output is close to that for a fixed-effects model when  $\hat{\tau}^2 \approx 0$ .

## 3.5 Example

As an example, we apply the **ipdforest** command to a dataset of four depression intervention studies. Data were provided by the authors of the studies, and we had complete information in terms of age, gender, exposure (control and intervention group membership), continuous outcome baseline, and endpoint values for 518 patients. Because the findings of the IPD meta-analysis had not been published when this article was being prepared, we used fake author names and generated random continuous and binary outcome variables for the purposes of this example while keeping the covariates at their actual values. We introduced correlation between baseline and endpoint scores and between-study variability, although the exact specification of the data generation is unimportant.

Using the semiartificial dataset, we perform a logistic IPD meta-analysis, followed by the **ipdforest** command.

```
. use ipdforest_example
. describe
Contains data from ipdforest_example.dta
                 518
  obs:
 vars:
                  17
                                               6 Feb 2012 11:35
              20,202
 size:
               storage
                       display
                                    value
variable name
                type
                        format
                                    label
                                                variable label
studyid
                byte
                        %22.0g
                                    stid
                                               Study identifier
                        %8.0g
patid
                                                Patient identifier
                 int
group
                byte
                        %20.0g
                                    grplbl
                                                Intervention/control group
                                    sexlbl
sex
                byte
                        %10.0g
                                               Gender
                float
                       %10.0g
                                               Age in years
age
                        %9.0g
depB
                byte
                                               Binary outcome, endpoint
depBbas
                byte
                        %9.0g
                                               Binary outcome, baseline
                byte
                        %9.0g
depBbas1
                                               Bin outcome baseline, trial 1
depBbas2
                        %9.0g
                byte
                                               Bin outcome baseline, trial 2
depBbas5
                        %9.0g
                byte
                                               Bin outcome baseline, trial 5
depBbas9
                        %9.0g
                byte
                                               Bin outcome baseline, trial 9
depC
                float
                       %9.0g
                                               Continuous outcome, endpoint
depCbas
                float
                       %9.0g
                                               Continuous outcome, baseline
depCbas1
                float
                       %9.0g
                                               Cont outcome baseline, trial 1
depCbas2
                       %9.0g
                                               Cont outcome baseline, trial 2
                float
depCbas5
                float %9.0g
                                               Cont outcome baseline, trial 5
depCbas9
                float %9.0g
                                               Cont outcome baseline, trial 9
```

```
Sorted by: studyid patid
```

We generate a centered age variable, interacted with the exposure variable in a mixed-effects logistic regression model. The model includes fixed study-specific intercepts and fixed study-specific effects for baseline and random treatment and age effects. The **ipdforest** command follows the regression model, requesting outcomes for both the main effect and the interaction.

```
. quietly summarize age
. quietly generate agec = age-r(mean)
. xtmelogit depB group agec sex i.studyid depBbas1 depBbas2 depBbas5 depBbas9
> i.group#c.agec || studyid:group agec, var nocons or
Refining starting values:
Iteration 0:
               \log likelihood = -347.40378
                                             (not concave)
Iteration 1:
               log likelihood = -336.07882
                                             (not concave)
Iteration 2:
               log likelihood = -329.28268
Performing gradient-based optimization:
Iteration 0:
               \log likelihood = -329.28268
                                              (not concave)
               \log likelihood = -326.79754
Iteration 1:
               log likelihood = -326.5689
Iteration 2:
Iteration 3:
               \log likelihood = -326.55747
Iteration 4:
               \log likelihood = -326.55747
Mixed-effects logistic regression
                                                 Number of obs
                                                                              518
Group variable: studyid
                                                 Number of groups
                                                                                4
                                                                               42
                                                 Obs per group: min =
                                                                           129.5
                                                                 avg =
                                                                              214
                                                                 max =
                                                                           42.06
Integration points =
                       7
                                                 Wald chi2(11)
                                                                     =
Log likelihood = -326.55747
                                                 Prob > chi2
                                                                     =
                                                                          0.0000
                                                 P>|z|
                                                            [95% Conf. Interval]
               Odds Ratio
                             Std. Err.
        depB
                                            z
                 1.840804
                             .3666167
                                          3.06
                                                 0.002
                                                            1.245894
                                                                         2.71978
       group
                  .9867902
                             .0119059
                                         -1.10
                                                 0.270
                                                            .9637288
                                                                        1.010403
        agec
                  .7117592
                             .1540753
                                         -1.57
                                                 0.116
                                                            .4656639
                                                                        1.087912
         sex
     studyid
                 1.050007
                             .5725515
                                          0.09
                                                 0.929
                                                            .3606168
                                                                        3.057302
          2
                                         -0.30
                                                 0.763
          5
                  .8014552
                             .5894511
                                                            .1896011
                                                                        3.387799
                                                            .4469621
          9
                 1.281413
                             .6886055
                                          0.46
                                                0.644
                                                                        3.673734
                                                 0.015
    depBbas1
                 3.152909
                             1.49528
                                          2.42
                                                            1.244587
                                                                        7.987251
                                          3.60
                                                 0.000
    depBbas2
                 4.480302
                            1.863908
                                                            1.982385
                                                                        10.12573
    depBbas5
                 2.387336
                             1.722993
                                          1.21
                                                 0.228
                                                            .5802064
                                                                        9.823007
    depBbas9
                 1.881203
                             .7086506
                                          1.68
                                                 0.093
                                                            .8990571
                                                                        3.936261
group#c.agec
                 1.011776
                             .0163748
                                          0.72
                                                 0.469
                                                            .9801858
                                                                        1.044385
          1
       _cons
                  .5533714
                             .2398341
                                         -1.37
                                                 0.172
                                                            .2366473
                                                                        1.293993
```

Random-effects Parameters	Estimate	Std. Err.	[95% Conf. I	interval]
studyid: Independent var(group) var(agec)		2.03e-11 4.41e-11	0 0	•

LR test vs. logistic regression: chi2(2) = 0.00 Prob > chi2 = 1.0000 Note: LR test is conservative and provided only for reference. . ipdforest group, fe(sex) re(agec) ia(agec) or

One-stage meta-analysis results using xtmelogit (ML method) and ipdforest Main effect (group)  $% \left( \left( x,y\right) \right) =\left( x,y\right) \right) =\left( x,y\right) +\left( x,y\right) \right) =\left( x,y\right) +\left( x,y\right) +\left( x,y\right) +\left( x,y\right) +\left( x,y\right) +\left( x,y\right) \right) +\left( x,y\right) +\left($ 

Study	Effect	[95% Conf.	Interval]	% Weight
Hart 2005	2.118	0.942	4.765	19.88
Richards 2004	2.722	1.336	5.545	30.69
Silva 2008	2.690	0.748	9.676	8.11
Kompany 2009	1.895	0.969	3.707	41.31
Overall effect	1.841	1.246	2.720	100.00

One-stage meta-analysis results using xtmelogit (ML method) and ipdforest Interaction effect (group x agec)

Study	Effect	[95% Conf.	Interval]	% Weight
Hart 2005	0.972	0.901	1.049	19.88
Richards 2004	0.995	0.937	1.055	30.69
Silva 2008	0.987	0.888	1.098	8.11
Kompany 2009	1.077	1.015	1.144	41.31
Overall effect	1.012	0.980	1.044	100.00

#### Heterogeneity Measures

	value	[95% Conf.	Interval]
I (%) H	•	0.000	
tau est	0.000	0.000	•

Maximum likelihood converged successfully in this example, and the between-study variance estimate  $\hat{\tau}^2$  was practically 0. Note that the intercept for the reference study (studyid = 1) was estimated in \_cons. The reported coefficients under studyid are the differences in intercept compared with the first study.  $I^2$  and  $H_M^2$  could not be estimated because residual variability is not reported under xtmelogit. The overall treatment effect was significant at the 95% level, but the overall effect for the interaction of treatment and age was not. The forest plots created by ipdforest are displayed in figures 1 and 2.

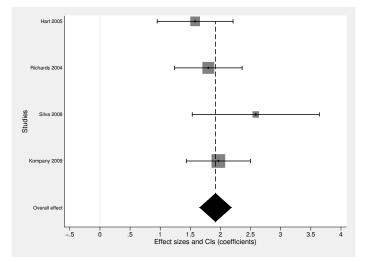


Figure 1. Main-effect IPD forest plot reporting odds ratios

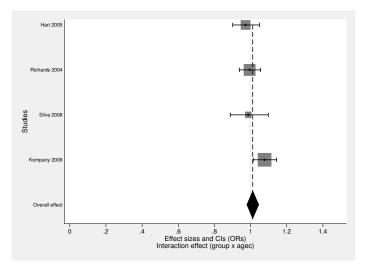


Figure 2. Interaction-effect IPD forest plot reporting odds ratios

# 4 Discussion

The aim of this article was to provide a practical guide for conducting one-stage IPD meta-analysis and to present ipdforest, a new forest-plot command. ipdforest aims to help meta-analysts better communicate their results through the familiar and distinctive forest plot—a graphical output not previously available in one-stage IPD meta-analysis software routines.

Although only binary or continuous exposure variables can be modeled, categorical exposures can also be investigated with the use of dummy variables and a focus on the comparison of interest through one of these. In addition, ipdforest is fully compatible with the estimates produced by the multiple-imputation estimation command mi estimate: xtmixed or mi estimate: xtmelogit.

Note that these commands were renamed in Stata 13: xtmixed to mixed and xtmelogit to meqrlogit. ipdforest is not yet compatible with the new commands, but users of version 13 can still use the older commands before calling ipdforest.

# 5 Acknowledgments

We would like to thank Isabel Canette, senior statistician at StataCorp, for her help with advanced aspects of xtmixed and xtmelogit and the anonymous reviewer whose comments and suggestions improved the command significantly. Evan Kontopantelis is on an NIHR School for Primary Care fellowship.

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