

Meta-analysis: Random-effects model

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nder the random-effects model, the true effect size is assumed to be different across studies. These studies are not identical (as in the case of the fixed-effect model), but they have enough in common to be included in the meta-analysis and synthesize their information.¹ For example, we are interested in identifying studies that examine the clinical effectiveness, such as bond failures of plasma vs halogen curing lights. We randomly sampled patients from different practices (also randomly selected). If plasma light is actually more effective than halogen, we expect the effect size (eg, risk ratio [RR] or odds ratio [OR]) to be similar but not identical across the practices. Because these practices can differ in their populations (ie, crowding severity, cooperation, age, and sex), implementation of interventions (ie, practitioner expertise, adhesives, and type of brackets used), outcomes (duration of follow-up), and settings (geographic area), among other reasons (known as covariates), the magnitude of the effect size is likely to vary across the studies; the effect size may be lower in practices with more expertise and older participants and vice versa. In practice, the existence of such covariates that affect the effect size is innate and leads to variations in the magnitude of the effect.

The random-effects model addresses the variations in the effect size across the studies beyond mere chance. According to this model, there is not a common effect size, but true study-specific effect sizes that are normally distributed about a mean (known as mean effect size) with variance that reflects the variation of these effect sizes. Our goal was to estimate the mean and the variance of this distribution. The summary effect size is actually the

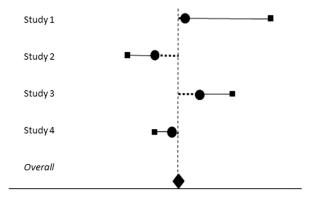


Fig 1. Forest plot of four fictional studies that illustrated the random-effects model.

mean effect size. Because in the random-effects model we study a sample of true effect sizes and not a common one, we use the plural (effects). Figure 1 presents a forest plot of 4 fictional studies under the random-effects model. In the random-effects model, each study provides a different effect size because of both within- and between-study variances (the latter implies the degree of dissimilarity of the included studies), unlike in the fixed-effect model in which only within-study variation is expected. Therefore, the weighting scheme of this model is based on 2 sources of variance in each study. The detailed steps to perform a random-effects meta-analysis are provided in the Supplementary Appendices 1 and 2.1

APPLICATION TO REAL DATA—BINARY OUTCOME

The application is based on the data set we used for the fixed-effect meta-analysis (Table 1). $^{2-7}$ The results under the random-effects model are almost the same (and slightly less precise) as those under the fixed-effect model (random-effects: RR = 0.92 with 95% confidence interval [CI] = 0.70-1.21 vs fixed-effect: RR = 0.90 with 95% CI = 0.69-1.19). This trivial difference was expected because the between-study variance was found to be trivial. The results of this worked example are illustrated in Figure 2. To simplify this

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Table I.	Dond	failures	hotwoon	nlacma	and	halagan	ouring	liah	+
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	Plasma			Halogen			
Study	Events	Nonevents	Sample size	Events	Nonevents	Sample size	
Cacciafesta, 2004 ²	21	279	300	12	288	300	
Manzo, 2004 ³	12	292	304	12	292	304	
Pettemerides, 2004 ⁴	12	164	176	13	163	176	
Russell, 2008 ⁵	22	305	327	31	296	327	
Sfondrini, 2004 ⁶	31	686	717	39	678	717	

Note: To perform a meta-analysis for binary data, it is necessary to have the sample size per group and the number of events.

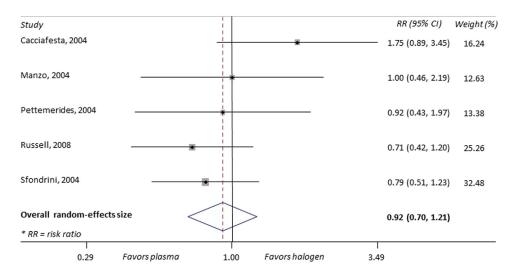


Fig 2. Forest plot of bond failures between plasma and halogen curing lights using the random-effects model—binary outcome.

example, clustering owing to multiple teeth bonded within patients was not considered during this analysis. Accounting for clustering would have increased the width of the 95% Cls, and hence the uncertainty of the estimates with no difference in the conclusions presented here.

APPLICATION TO REAL DATA—CONTINUOUS OUTCOME

For the random-effects meta-analysis on continuous data, we used the data set we applied for the fixed-effect meta-analysis (Table II). The results for standardized mean difference (SMD) under the random-effects model are almost the same (and less precise) as those under the fixed-effect model (random-effects: SMD = -0.14 with 95% CI = -0.47 to 0.19 vs fixed-effect: SMD = -0.13 with 95% CI = -0.33 to 0.07). This difference in the width of 95% CIs was expected because the

between-study variance was found to be equal to 0.08. The results of this worked example are illustrated in Figure 3.

Table II. Initial orthodontic alignment using Damon and conventional appliances

		Damon			Conventional			
Study	Mean	SD	Sample size	Mean	SD	Sample size		
Scott, 2008 ⁹	1.41	1.41	32	1.68	0.66	28		
Ong, 2010 ¹⁰	1.46	0.65	44	1.57	0.59	40		
Pandis, 2011	1.98	0.90	25	2.22	0.96	25		
Miles, 2006 ¹²	0.51	0.66	29	0.60	0.75	29		
Pandis, 2007 ¹³	1.81	0.71	27	1.41	0.63	27		
Wahab, 2012 ¹⁴	2.16	1.08	14	3.15	1.17	15		

Note: To perform a meta-analysis for continuous data, it is necessary to have the sample size, the mean, and the SD per group. *SD*, standard deviation.

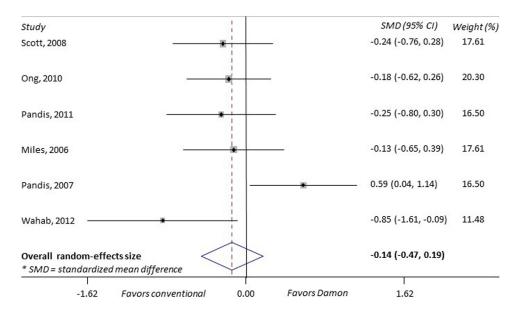


Fig 3. Forest plot of effectiveness of initial orthodontic alignment between Damon and conventional appliances using the random-effects model—continuous outcome.

SUPPLEMENTARY DATA

Supplementary data associated with this article can be found, in the online version, at https://doi.org/10.1016/j.ajodo.2019.10.007.

REFERENCES

- Borenstein M, Hedges LV, Higgins JP, Rothstein HR. A basic introduction to fixed-effect and random-effects models for meta-analysis. Res Synth Methods 2010;1:97-111.
- Cacciafesta V, Sfondrini MF, Scribante A. Plasma arc versus halogen light-curing of adhesive-precoated orthodontic brackets: a 12-month clinical study of bond failures. Am J Orthod Dentofacial Orthop 2004;126:194-9.
- 3. Manzo B, Liistro G, De Clerck H. Clinical trial comparing plasma arc and conventional halogen curing lights for orthodontic bonding. Am J Orthod Dentofacial Orthop 2004;125:30-5.
- Pettemerides AP, Sherriff M, Ireland AJ. An in vivo study to compare a plasma arc light and a conventional quartz halogen curing light in orthodontic bonding. Eur J Orthod 2004;26: 573-7.
- Russell JS, Littlewood SJ, Blance A, Mitchell L. The efficacy of a plasma arc light in orthodontic bonding: a randomized controlled clinical trial. J Orthod 2008;35:202-9.
- Sfondrini MF, Cacciafesta V, Scribante A, Klersy C. Plasma arc versus halogen light curing of orthodontic brackets: a 12-month clinical study of bond failures. Am J Orthod Dentofacial Orthop 2004;125:342-7.
- 7. Fleming PS, Eliades T, Katsaros C, Pandis N. Curing lights for orthodontic bonding: a systematic review and

- meta-analysis. Am J Orthod Dentofacial Orthop 2013;143: 592-103.
- Pandis N, Walsh T, Polychronopoulou A, Eliades T. Cluster randomized clinical trials in orthodontics: design, analysis and reporting issues. Eur J Orthod 2013;35:669-75.
- Scott P, DiBiase AT, Sherriff M, Cobourne MT. Alignment efficiency of Damon3 self-ligating and conventional orthodontic bracket systems: a randomized clinical trial. Am J Orthod Dentofacial Orthop 2008;134:470.e1–8.
- Ong E, McCallum H, Griffin MP, Ho C. Efficiency of self-ligating vs conventionally ligated brackets during initial alignment. Am J Orthod Dentofacial Orthop 2010;138:138.e1–7.
- 11. Pandis N, Polychronopoulou A, Katsaros C, Eliades T. Comparative assessment of conventional and self-ligating appliances on the effect of mandibular intermolar distance in adolescent nonextraction patients: a single-center randomized controlled trial. Am J Orthod Dentofacial Orthop 2011;140:e99-105.
- 12. Miles PG, Weyant RJ, Rustveld L. A clinical trial of Damon 2 vs conventional twin brackets during initial alignment. Angle Orthod 2006;76:480-5.
- Pandis N, Polychronopoulou A, Eliades T. Self-ligating vs conventional brackets in the treatment of mandibular crowding: a prospective clinical trial of treatment duration and dental effects.
 Am J Orthod Dentofacial Orthop 2007;132:208-15.
- Wahab RMA, Idris H, Yacob H, Ariffin SHZ. Comparison of self- and conventional-ligating brackets in the alignment stage. Eur J Orthod 2011;34:176-81.
- Pandis N, Fleming PS, Spineli LM, Salanti G. Initial orthodontic alignment effectiveness with self-ligating and conventional appliances: a network meta-analysis in practice. Am J Orthod Dentofacial Orthop 2014;145:S152-63.