Discrepancies in Outcome Reporting Exist Between Protocols and Published Oral Health Cochrane Systematic Reviews

Nikolaos Pandis¹,²*, Padhraig S. Fleming³, Helen Worthington⁴, Kerry Dwan⁵, Georgia Salanti¹

¹ Department of Hygiene and Epidemiology, University of Ioannina School of Medicine, Ioannina, Greece,
² Department of Orthodontics and Dentofacial Orthopedics, Dental School/Medical Faculty, University of Bern, Bern, Switzerland,
³ Barts and The London School of Medicine and Dentistry, Queen Mary University of London, London, United Kingdom,
⁴ Cochrane Oral Health Group, School of Dentistry, The University of Manchester, Manchester, United Kingdom,
⁵ Institute of Translational Medicine, Department of Biostatistics, University of Liverpool, Liverpool, United Kingdom

* npandis@yahoo.com

Abstract

Objectives
To assess discrepancies in the analyzed outcomes between protocols and published reviews within Cochrane oral health systematic reviews (COHG) on the Cochrane Database of Systematic Reviews (CDSR).

Study Design and Setting
All COHG systematic reviews on the CDSR and the corresponding protocols were retrieved in November 2014 and information on the reported outcomes was recorded. Data was collected at the systematic review level by two reviewers independently.

Results
One hundred and fifty two reviews were included. In relation to primary outcomes, 11.2% were downgraded to secondary outcomes, 9.9% were omitted altogether in the final publication and new primary outcomes were identified in 18.4% of publications. For secondary outcomes, 2% were upgraded to primary, 12.5% were omitted and 30.9% were newly introduced in the publication. Overall, 45.4% of reviews had at least one discrepancy when compared to the protocol; these were reported in 14.5% reviews. The number of review updates appears to be associated with discrepancies between final review and protocol (OR: 3.18, 95% CI: 1.77, 5.74, p<0.001). The risk of reporting significant results was lower for both downgraded outcomes [RR: 0.52, 95% CI: 0.17, 1.58, p = 0.24] and upgraded or newly introduced outcomes [RR: 0.77, 95% CI: 0.36, 1.64, p = 0.50] compared to outcomes with no discrepancies. The risk of reporting significant results was higher for upgraded or newly introduced outcomes compared to downgraded outcomes (RR = 1.19, 95% CI: 0.65, 2.16, p = 0.57). None of the comparisons reached statistical significance.
Conclusion
While no evidence of selective outcome reporting was found in this study, based on the present analysis of SRs published within COHG systematic reviews, discrepancies between outcomes in pre-published protocols and final reviews continue to be common. Solutions such as the use of standardized outcomes to reduce the prevalence of this issue may need to be explored.

Introduction
Systematic reviews (SRs) for interventions provide the basis for evaluating the body of evidence and by extension are uniquely influential in leading to healthcare recommendations [1,2]. However, inappropriate practices such as selective inclusion of trials, selective reporting of outcomes and results-driven reporting can bias estimates of treatment effects culminating in compromised patient care, healthcare decisions and service configuration [3,4]. Selective outcome reporting involving preferential reporting of specific data or outcomes within a study is a recognized problem and has been investigated in respect of randomized controlled trials (RCTs), in particular [5–9].

While research on selective reporting has focused both on RCTs and non-randomized studies [10,11], similar issues may arise within SRs [12]. At the systematic review level, selective reporting may develop for a variety of reasons, for example, due to the use of multiple measurement scales, outcomes or time points [13,14] and selective inclusion of specific outcomes. Arbitrary inclusion of outcomes based on post hoc results-driven decisions may bias the conclusions from subsequent syntheses [13,15].

A recent systematic review [15] has highlighted among other issues the paucity of information on selective reporting based on comparisons between review protocols and final systematic review reports. The limited number of meta-epidemiological analyses identified were restricted to cystic fibrosis reviews on the Cochrane Database of Systematic Reviews (CDSR) prior to 2010 [16], or involved analysis of selected issues within the CDSR over three time periods between 2000 and 2008 [17–19].

Historically, with the notable exception of the CDSR, systematic reviews have lacked pre-published protocols. This has changed following the relatively recent introduction of the PROSPERO database, and publication of SR protocols in isolation in a dedicated journal (www.systematicreviewsjournal.com)[20], with reporting on the existence of a pre-published protocol also encouraged within the PRISMA guidelines [2]. To our knowledge, while the existence and impact of outcome reporting bias within dental SRs has been acknowledged [21], there are no reports addressing selective reporting and discrepancies between systematic review protocols and final reports within the oral health field. Therefore, the aim of the present study was to explore the prevalence and nature of selective reporting and factors associated with selective reporting within SRs published by the Cochrane Oral Health Group (COHG) in the CDSR by analyzing discrepancies between systematic review protocols and final reports.

Methods
All systematic reviews in the Cochrane Oral Health Group (COHG) published in the CDSR until November, 2014 were screened for final reports and pre-published protocols. The first author (NP) contacted the COHG in order to retrieve review protocols that were not available
in the Cochrane library. Reviews where the protocol could not be found as well as those reviews published in duplicate were to be excluded from further analysis.

Data were independently extracted and entered on pre-piloted standardized forms for the eligible studies. Initial calibration was performed between the two researchers (NP, PSF) on 10 articles. Disagreements were resolved by discussion or, if necessary, with adjudication by a third reviewer (KD). Information obtained included the number and type of primary and secondary outcomes both within the protocol and in the final publication. If a review did not distinguish between primary and secondary outcomes [unlabelled outcomes], the first three outcomes listed were taken to be the primary outcomes and the rest considered as secondary outcomes. Specific details as to whether outcomes present in the protocol were omitted, upgraded or downgraded as well as inclusion of outcomes in the final report that were absent from the protocol were recorded. Data relating to year of publication of the protocol and the final report, and any previous versions of the review, number of authors, geographical location of the corresponding author, whether the collaboration involved a single or multiple centers and subject area were also extracted. In addition, information concerning statistical significance was extracted for one primary outcome from the reviews with applicable meta-analyses. The primary review comparison was assumed for each review[19] according to the following hierarchy by selecting that which met the first of the following criteria: (1) an intervention comparison described in the protocol as the primary review comparison; (2) the first intervention comparison mentioned in the objectives of the protocol; (3) an intervention comparison described in the review as the primary review comparison; (4) the first intervention comparison mentioned in the objectives of the review; (5) the intervention comparison used in the first meta-analysis presented in the review.

Data Analysis

Descriptive statistics were calculated for all included articles related to the following variables: geographical representation, number of research centers, number of review updates, number of authors and year of publication. The year of publication was converted to a binary variable [<2008, ≥2008, as from 2008 Cochrane included a requirement to declare changes between protocol and review]. Cross-tabulations were undertaken to investigate associations between the presence or absence of discrepancies and review characteristics. A logistic model was also fitted in order to assess and quantify any association of discrepancies between protocol and final review and review characteristics. Finally, upgrades, downgrades, new introductions and omissions between protocol entries and final reports for all outcomes listed were tabulated. Risk ratios were calculated in order to examine the relationship between possible discrepancies in the primary outcomes and statistical significance. All statistical analyses were conducted with Stata™ version 13.1 software (Stata Corporation, College Station, Texas, USA).

Results

Initially 172 titles were identified in the COHG database of which 19 reviews were excluded as duplicates [same title and DOI] and one diagnostic review was excluded (Fig 1). From the remaining 152 reviews 116 (76.3%) had accessible protocols next to the review under the “Protocol and previous versions” section on the Cochrane Library. The COHG provided the protocols for 36 reviews for which the protocols were not available in the Cochrane library.

A variety of conditions, interventions and outcomes were considered with the highest number of reviews dealing with dental caries, orthodontics, periodontics and implantology (S1 Table). The largest proportion of reviews originated in the UK according to the details of the corresponding authors (S2 Table).
The median number of labelled primary outcomes was 2 (range: 0–11) and secondary outcomes was 4 (range: 0–36) both in the protocols and published reviews. Overall, 51 protocols (33.6%) and 34 final reviews (22.4%) did not distinguish between primary and secondary outcomes. This was less common among reviews published after 2008 compared to those published prior to 2008 (OR: 0.33, 95% CI: 0.14, 0.75, \(p = 0.01\)) but no important improvement was observed at protocol level for outcome distinction after 2008 compared to the previous period (OR: 0.54, 95% CI: 0.25, 1.16, \(p = 0.11\)). The distribution of individual discrepancies between protocols and reviews per geographical area, single or multicenter collaboration, number of review updates, number of authors and publication period is presented in Table 1.

According to the multivariable logistic model (Table 2) the number of review updates appears to be associated with discrepancies between final review and protocol (OR: 3.18, 95% CI: 1.77, 5.74, \(p < 0.001\)). Geographical area, single or multicenter collaboration, review publication year and number of authors do not appear to be important predictors of discrepancies between protocols and final reports.

The detailed analysis of discrepancies (Table 3) revealed that the primary outcome was not stated in 3.9% (6/152) of protocols but stated in all final reports. In 30.1% (49/152) of the reviews the primary outcomes were not consistent in the protocol and the report while secondary outcomes were not the same in 43.4% of the reviews (66/152). In 11.2% (17/152) of the reviews the primary outcomes were downgraded to secondary, in 9.9% of the reviews (15/152) they were omitted and in 18.4% (28/152) new primary outcomes were introduced. In four reviews out of 152 (2.6%) a change in the definition was identified. In 2% (3/152) of the reviews secondary outcomes were upgraded to primary, 12.5% (19/152) were omitted and in 30.9% (47/152) new secondary outcomes were introduced. No outcome definition change was identified for secondary outcomes. Overall, 45.4% (69/152) of the reviews presented at least one discrepancy but only in 14.5% (10/69), all published after 2008, were discrepancies reported in the...
Discrepancies between Protocols and Published Systematic Reviews

Table 1. Distribution of any discrepancies between protocols and reviews per geographical area, single or multicenter collaboration, number of review updates, number of authors and review publication period.

| Continent of origin of corresponding author | Discrepancy | p-value |
|--------------------------------------------|-------------|---------|
|                                            | Total       | no      | no      | yes     | yes     | 0.09* |
|                                            | No.         | No.     | % No.   | No.     | % No.   |
| Middle East & Africa                       | 10          | 7       | 70      | 3       | 30      |
| Americas                                   | 15          | 12      | 80      | 3       | 20      |
| Asia                                       | 11          | 7       | 64      | 4       | 36      |
| Europe                                     | 116         | 57      | 49      | 59      | 51      |
| Total                                      | 152         | 79      | 52      | 73      | 48      |

Collaboration between centers

| No. | Yes | 0.41* |
|-----|-----|-------|
| 4   | 148 | 3     |
| 3   | 1   | 25    |

Number of review updates

| No updates | 1 update | ≥2 updates | 0.001** |
|------------|----------|------------|---------|
| 104        | 29       | 19         | <0.001**|
| 69         | 12       | 2          |         |
| 66         | 41       | 11         |         |
| 35         | 17       | 17         |         |
| 34         | 59       | 89         |         |

Number of authors

| 2–3         | 4–5       | >6         | 0.98** |
|-------------|-----------|------------|--------|
| 21          | 73        | 58         | 48     |
| 11          | 40        | 32         | 45     |
| 52          | 55        | 55         | 45     |
| 10          | 33        | 26         | 45     |
| 48          | 45        | 45         |        |

Review publication period

| <2008       | ≥2008     | 0.10** |
|-------------|-----------|--------|
| 36          | 116       | 55     |
| 24          | 59        | 69     |
| 67          | 51        | 45     |
| 12          | 57        | 49     |
| 33          | 49        |        |

Total

| Total       | 152       | 83       | 55     | 69     | 45     |

* Fisher’s exact test
** Chi² test

doi:10.1371/journal.pone.0137667.0001

text (S3 Table). It is important to point out that not all declared changes were explained in detail or justified.

Table 2. Adjusted odds ratios and associated 95% confidence intervals for the association between any discrepancies (protocol vs final report) and review characteristics.

| Characteristic                  | OR   | 95% CI       | p-value |
|---------------------------------|------|--------------|---------|
| Number of authors [per unit]    | 1.10 | 0.90, 1.34   | 0.34    |
| Geography region
| Americas, reference             | -    | -            | -       |
| Middle East & Africa            | 1.95 | 0.29, 13.34  | 0.50    |
| Asia                            | 2.15 | 0.35, 13.33  | 0.41    |
| Europe                          | 3.09 | 0.78, 12.29  | 0.11    |
| Number of updates [per unit]    | 3.18 | 1.77, 5.74   | <0.001  |
| Review publication year
| <2008, reference                | -    | -            | -       |
| ≥2008                           | 1.33 | 0.54, 3.29   | 0.54    |
| Collaboration between centers   | No, reference                    | -    | -            | -       |
| Yes                             | 2.55 | 0.16, 41.65  | 0.51    |

doi:10.1371/journal.pone.0137667.0002
Of the 152 reviews assessed, 89 did not have a meta-analysis or the meta-analysis did not relate to the primary outcomes. Within the remaining reviews the distribution of discrepancies for the primary outcome by statistical significance is shown in Table 4. The risk of reporting significant results was lower for downgraded outcomes [RR: 0.52, 95% CI: 0.17, 1.58, p = 0.24] and upgraded or newly introduced outcomes [RR: 0.77, 95% CI: 0.36, 1.64, p = 0.50]. The risk of a significant result was higher for upgraded/new outcomes (RR = 1.19, 95% CI: 0.65, 2.16, p = 0.57) compared to outcome downgrades. None of the comparisons reached statistical significance.

### Discussion

#### Main findings

This meta-epidemiological study attempted to identify discrepancies between 152 COHG systematic reviews and protocols in order to shed some light on selective reporting in oral health.

### Table 3. Tabulation of type of discrepancy between protocols and reviews for primary and secondary outcomes at the review level.

| Type of discrepancy                                                                 | n   | %   |
|-------------------------------------------------------------------------------------|-----|-----|
| Primary outcome variable not stated in the protocol                                | 6/152 | 3.9 |
| Primary outcome variable not stated in the published report                         | 0/152 | 0.0 |
| All primary outcome(s) stated in the protocol is/are not the same as in the published report | 47/152 | 30.9 |
| One or several primary outcome(s) stated in the protocol is downgraded to secondary in the published report | 17/152 | 11.2 |
| One or several primary outcome(s) stated in the protocol is/are omitted from the published report | 15/152 | 9.9 |
| One or several new primary outcome(s) that was/were not stated in the protocol is included in the published report | 28/152 | 18.4 |
| The definition of one or several primary outcome(s) was different in the protocol compared to the published report | 4/152 | 2.6 |
| All secondary outcomes stated in the protocol are not the same as in the published report | 66/152 | 43.4 |
| One or several non-primary outcome(s) in the protocol is/are changed to primary in the published report | 3/152 | 2.0 |
| One or several secondary outcome(s) stated in the protocol is/are omitted from the published | 19/152 | 12.5 |
| One or several new secondary outcome(s) that was/were not stated in the protocol is/are included in the published report | 47/152 | 30.9 |
| The definition of one or several secondary outcome(s) was different in the protocol compared to the published report | 0/152 | 0.0 |

Of the 152 reviews assessed, 89 did not have a meta-analysis or the meta-analysis did not relate to the primary outcomes. Within the remaining reviews the distribution of discrepancies for the primary outcome by statistical significance is shown in Table 4. The risk of reporting significant results was lower for downgraded outcomes [RR: 0.52, 95% CI: 0.17, 1.58, p = 0.24] and upgraded or newly introduced outcomes [RR: 0.77, 95% CI: 0.36, 1.64, p = 0.50]. The risk of a significant result was higher for upgraded/new outcomes (RR = 1.19, 95% CI: 0.65, 2.16, p = 0.57) compared to outcome downgrades. None of the comparisons reached statistical significance.

### Table 4. Tabulation of counts, risk ratios and associated 95% confidence intervals and p-values per type of discrepancy and statistical significance for primary outcomes only.

|                        | Total | Significant (<0.05) | Non-significant (>0.05) | Risk Ratio (95% CI) | p-value |
|------------------------|-------|---------------------|-------------------------|---------------------|---------|
| No discrepancy         | 24    | 16                  | 8                       | Reference           | -       |
| Downgrade              | 9     | 4                   | 5                       | 0.52 (0.17, 1.58)   | 0.24    |
| Upgrade/Inclusion      | 16    | 9                   | 7                       | 0.77 (0.36, 1.64)   | 0.50    |
|                         | Total | Significant (<0.05) | Non-significant (>0.05) | Risk Ratio (95% CI) | p-value |
| Downgrade              | 9     | 4                   | 5                       | Reference           | -       |
| Upgrade/Inclusion      | 16    | 9                   | 7                       | 1.19 (0.65, 2.16)   | 0.57    |
systematic reviews. The reviews covered a wide range of topics and were produced by authors located predominantly in the UK. Similar reviews have been undertaken within Cystic Fibrosis reviews on the CDSR [16] or generally within specific issues of the CDSR covering a range of review groups [17–19]. The most recent of the latter reviews focused on a period up to 2010 [16] identifying a discrepancy rate of up to 39%, while an earlier review alluded to a discrepancy rate of 22% [17]. Our findings spanning a 16-year period indicate that within SRs in the COHG discrepancies continue to be highly prevalent, with a prevalence of 45% for primary outcomes. The risk for discrepancy increased with increasing number of review updates. This is intuitive as additional outcomes can become more or less important over time with, for example, increasing emphasis on important outcomes in recent years. Moreover, new reviewers may differ from previous reviewers in relation to the priority with which they assign to certain outcomes. The risk of statistically significant results was lower for both outcome downgrades and upgrades/new inclusion compared to outcomes with no discrepancy. The finding for upgrades/new inclusions was opposite to what was expected. The risk of statistically significant results, consistent with other studies, was higher for outcome upgrades/new inclusions compared to outcome downgrades. None of those findings reach statistical significance however, as data was thin.

Some of these problems with discrepancies between protocols and reviews will be addressed when the COHG completes research being undertaken on Core Outcome Measures in Effectiveness Trials (COMET; http://www.comet-initiative.org) which will provide a list of core outcomes to be included in reviews being undertaken in certain areas of oral health.

Limitations and strengths

A large number of reviews were captured relative to published previous assessments, which have ranged from 46 to 288 SRs [16,17]. Articles were identified by just one author in the present study; however, as just a single database was searched with articles readily accessible, the risk of selection bias and inappropriate omission of relevant articles is very low. We are, therefore, confident that the present findings are a true reflection of the discrepancy rate both for primary and secondary outcomes with the Oral Health Group of the CDSR. The Oral Health Group relates to reviews published within the dentistry and oral and maxillofacial surgery; previous meta-epidemiological assessments have confirmed that dental SRs are equally susceptible to methodological and reporting weaknesses as other biomedical areas [22–25]. It is, therefore, likely that the findings from the present review are representative of the CDSR more broadly.

Findings in context

A large number of discrepancies were identified between protocols and final reports for both primary and secondary outcomes; these included introduction of new outcomes, upgrades, downgrades, omissions and definition changes. Since 2008, the Cochrane Collaboration requires that published reviews disclose discrepancies between protocols and reports. In this study a low percentage (14.5%) of outcome discrepancies were declared. This figure is in keeping with similar research with Kirkham et al. reporting acknowledgment of post hoc changes in just 6% of 64 reviews found to have discrepancies [17]. However, in a more recent review [16] discrepancies were mentioned in 39%, albeit based on a smaller sample of 46 reviews overall. These figures, however, may indicate a lack of awareness among review authors and failure of reviewers to cross-reference final submissions with pre-published protocols. While there is ample evidence that SRs published on the CDSR are of higher methodological quality than those published elsewhere [26–28] the stipulation of disclosure of discrepancies within the
CDSR does not appear to have had the desired effect either within the COHG or more generally.

In the present subset, no statistically significant association between alteration in outcomes (upgrade from secondary to primary, introduction of new outcomes, downgrades) and statistically significant of the meta-analysis results was found. The risk for reporting significant results was higher for upgraded or newly introduced outcomes compared to downgraded outcomes, although a non-statistically significant finding. Kirkham et al. [19] found an increased risk of a significant result with discrepancies including new inclusion or upgrades but not with downgrades. The discordance in relation to upgrades/new inclusion of outcomes between these studies may be attributed to the relatively small number of included meta-analyses in the present study but may also relate to the type of relevant outcomes.

**Implications of results**

The continued stubbornly high outcome discrepancy rate within the CDSR suggests that the prerequisite of delineating changes between protocol and submitted review is ineffective. However, external screening by the Cochrane Editorial Unit from 2013 has helped to ensure these changes are documented and justified in new reviews published since September 2013.

An alternative may be to consider a streamlined approach incorporating checks ensuring that outcomes in the review correspond with those delineated in the protocols. This approach could be implemented at the editorial level with proprietary software. For example, outcomes from the protocol may be automatically transferred to the review. Where authors wish to make changes in the review those changes would only be permitted if a declaration for changes field was completed. The editor could then be alerted of the outcome changes between protocol and review prompting the need for approval and registering the change. In order to improve transparency consideration could be given to publication of formal protocol amendments within the CDSR, although this would require external approval and may lead to a more protracted publication process.

The importance of setting core outcomes common to individual trials and reviews in order to enhance usability and transparency of reports and to reduce variability in outcome specification [29] can also be considered as an important step in reducing discrepancies [30,31]. However, the introduction of standardised core outcome sets may, initially at least, result in more discrepancies between reviews and protocols (particularly in review updates) as review authors adjust their outcomes to reflect consensus core outcome sets. In time, however, it is likely that as core outcome sets become more established it is likely that the prevalence of selective reporting and outcome alterations would diminish.

In terms of predictors, it appears differences between protocols and published reviews were more likely with increasing numbers of review updates. This finding is intuitive and may reflect the fact that reviews may span periods in excess of a decade and may, therefore, involve different author groups. Moreover, there are instances of multiple reviews being derived from the same protocol. The correct protocols for these “split” reviews were provided by the COHG group office and were clearly identified. It may, however, be sensible to consider more regular update of SR protocols within the CDSR after a defined time period to mitigate this.

In conclusion, the requirement to declare outcome changes between protocols and reviews which was implemented in 2008 is certainly a positive, there is further evidence that discrepancies still exist based on this analysis of SRs published within the Oral Health Group since its establishment. Alternative approaches to reduce the prevalence of this issue including the use of Core Outcome Sets need to be explored.
Supporting Information

S1 File. Dataset.
(XLSX)

S1 Table. Distribution of discrepancies between protocols and reviews per subject area.
(DOCX)

S2 Table. Distribution of discrepancies between protocols and reviews per country of origin.
(DOCX)

S3 Table. Explanation provided by authors for outcome changes between protocol and review.
(DOCX)

Acknowledgments

We would like to thank Jo Weldon from the COHG for providing the missing protocols.

Author Contributions

Conceived and designed the experiments: NP HW KD GS. Performed the experiments: NP PSF. Analyzed the data: NP KD GS. Contributed reagents/materials/analysis tools: N/A. Wrote the paper: NP PSF HW KD GS.

References

1. Higgins JPT, Altman DG, Sterne JAC (editors). Chapter 8: Assessing risk of bias in included studies. In: Higgins JPT, Green S (editors). Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0 (updated March 2011). The Cochrane Collaboration, 2011. Available: www.cochrane-handbook.org.
2. Moher D, Liberati A, Tetzlaff J, Altman DG, PRISMA Group. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. J Clin Epidemiol. 2009; 62: 1006–1012. doi:10.1016/j.jclinepi.2009.06.005 PMID: 19631508
3. Hutton JL, Williamson PR. Bias in meta-analysis due to outcome variable selection within studies. Applied Statistics 2000; 49(3):359–70.
4. Song F, Parekh S, Hooper L, Loke YK, Ryder J, Sutton AJ, et al. Dissemination and publication of research findings: an updated review of related biases. Health Technology Assessment 2010; 14(8):i, ix–xi, 1–193. doi: 10.3310/hta14080 PMID: 20181324
5. Dwan K, Altman DG, Arnaiz JA, Bloom J, Chan A-W, Cronin E, et al. Systematic review of the empirical evidence of study publication bias and outcome reporting bias. PloS One. 2008; 3: e3081. doi: 10.1371/journal.pone.0003081 PMID: 18769481
6. Dwan K, Altman DG, Clarke M, Gamble C, Higgins JPT, Sterne JAC, et al. Evidence for the selective reporting of analyses and discrepancies in clinical trials: a systematic review of cohort studies of clinical trials. PLoS Med. 2011; 11: e1001666. doi: 10.1371/journal.pmed.1001666 PMID: 24959719
7. Kirkham JJ, Dwan KM, Altman DG, Gamble C, Dodd S, Smyth R, et al. The impact of outcome reporting bias in randomised controlled trials on a cohort of systematic reviews. BMJ. 2010; 340: c365. doi: 10.1136/bmj.c365 PMID: 20156912
8. Chan A-W, Altman DG. Identifying outcome reporting bias in randomised trials on PubMed: review of publications and survey of authors. BMJ. 2005; 330: 753. doi: 10.1136/bmj.38356.424606.8F PMID: 15681569
9. Chan A-W, Hróbjartsson A, Haahr MT, Gotzsche PC, Altman DG. Empirical evidence for selective reporting of outcomes in randomized trials: comparison of protocols to published articles. JAMA J Am Med Assoc. 2004; 291: 2457–2465. doi: 10.1001/jama.291.20.2457
10. Booth A, Clarke M, Dooley G, Ghersi D, Moher D, Petticrew M, et al. The nuts and bolts of PROSPERO: an international prospective register of systematic reviews. Syst Rev. 2012; 1: 2. doi: 10.1186/2046-4053-1-2 PMID: 22587842
11. Norris SL, Moher D, Reeves BC, Shea B, Loke Y, Garner S, et al. Issues relating to selective reporting when including non-randomized studies in systematic reviews on the effects of healthcare interventions. Research Synthesis Methods 2013; 4(1):36–47. doi:10.1002/jrsm.1062 PMID: 26053538

12. Page MJ, McKenzie JE, Forbes A. Many scenarios exist for selective inclusion and reporting of results in randomized trials and systematic reviews. J Clin Epidemiol. 2013; 66: 524–537. doi:10.1016/j.jclinepi.2012.10.010 PMID: 23377885

13. Tendal B, Higgins JPT, Jüni P, Hróbjartsson A, Trelle S, Nüesch E, et al. Disagreements in meta-analyses using outcomes measured on continuous or rating scales: observer agreement study. BMJ. 2009; 339: b3128. doi:10.1136/bmj.b3128 PMID: 19679616

14. Bender R, Bunce C, Clarke M, Gates S, Lange S, Pace NL, et al. Attention should be given to multiplicity issues in systematic reviews. J Clin Epidemiol. 2008; 61: 857–865. doi:10.1016/j.jclinepi.2008.03.004 PMID: 18687287

15. Page MJ, McKenzie JE, Kirkham J, Dwan K, Kramer S, Green S, et al. Bias due to selective inclusion and reporting of outcomes and analyses in systematic reviews of randomised trials of healthcare interventions. Cochrane Database Syst Rev. 2014; 10: MR000035. doi:10.1002/14651858.MR000035.pub2

16. Dwan K, Kirkham JJ, Williamson PR, Gamble C. Selective reporting of outcomes in randomised controlled trials in systematic reviews of cystic fibrosis. BMJ Open. 2013; 3: doi:10.1136/bmjopen-2013-002709

17. Silagy CA, Middleton P, Hopewell S. Publishing protocols of systematic reviews: comparison of what was done to what was planned. JAMA. 2002; 287: 2831–2834. PMID: 12038926

18. Parmelli E, Liberati A, D’Amico R. Reporting of outcomes in systematic reviews: comparison of protocols and published systematic reviews (abstract). XV Cochrane Colloquium; 2007 Oct 23–27; Säo Paulo, Brazil. 2007: 118–9.

19. Kirkham JJ, Altman DG, Williamson PR. Bias due to changes in specified outcomes during the systematic review process. PloS One. 2010; 5: e9810. doi:10.1371/journal.pone.0009810 PMID: 20339557

20. www.systematicreviewsjournal.com.

21. Millett D. Bias in systematic reviews? J Orthod. 2011; 38: 158–160. doi:10.1179/14653121141407 PMID: 21875988

22. Seehra J, Fleming PS, Polychronopoulou A, Pandis N. Reporting completeness of abstracts of systematic reviews published in leading dental specialty journals. Eur J Oral Sci. 2013; 121: 57–62. doi:10.1111/eos.12027 PMID: 23489893

23. Kiriakou J, Pandis N, Fleming PS, Madianos P, Polychronopoulou A. Reporting quality of systematic review abstracts in leading oral implantology journals. J Dent. 2013; 41: 1181–1187. doi:10.1016/j.jdent.2013.09.006 PMID: 24075952

24. Fleming PS, Seehra J, Polychronopoulou A, Fedorowicz Z, Pandis N. A PRISMA assessment of the reporting quality of systematic reviews in orthodontics. Angle Orthod. 2013; 83: 158–163. doi:10.2319/032612-251.1 PMID: 22720835

25. Papageorgiou SN, Papadopoulos MA, Athanasiou AE. Reporting characteristics of meta-analyses in orthodontics: methodological assessment and statistical recommendations. Eur J Orthod. 2014; 36: 74–85. doi:10.1093/ejor/cjt008 PMID: 23493385

26. Fleming PS, Seehra J, Polychronopoulou A, Fedorowicz Z, Pandis N. Cochrane and non-Cochrane systematic reviews in leading orthodontic journals: a quality paradigm? Eur J Orthod. 2013; 35: 244–248. doi:10.1093/ejor/jct016 PMID: 22510325

27. Remschmidt C, Wichmann O, Harder T. Methodological quality of systematic reviews on influenza vaccination. Vaccine. 2014; 32: 1678–1684. doi:10.1016/j.vaccine.2014.01.060 PMID: 24513008

28. Moher D, Tetzlaff J, Tricco AC, Sampson M, Altman DG. Epidemiology and reporting characteristics of systematic reviews. PLoS Med. 2007; 4: e78. doi:10.1371/journal.pmed.0040078 PMID: 17388659

29. Saldanha LJ, Dickersin K, Wang X, Li T. Outcomes in Cochrane systematic reviews addressing four common eye conditions: an evaluation of completeness and comparability. PloS One. 2014; 9: e109400. doi:10.1371/journal.pone.0109400 PMID: 25329377

30. Smith V, Clarke M, Williamson P, Gargon E. Survey of new 2007 and 2011 Cochrane reviews found 37% of prespecified outcomes not reported. J Clin Epidemiol. 2015; 68: 237–245. doi:10.1016/j.jclinepi.2014.09.022 PMID: 25455837

31. Wuytack F, Smith V, Clarke M, Williamson P, Gargon E. Towards core outcome set (COS) development: a follow-up descriptive survey of outcomes in Cochrane reviews. Syst Rev. 2015; 4: 73. doi:10.1186/s13643-015-0060-0 PMID: 25987294