

1 **Factors predicting publication of Cochrane reviews**

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21**Word count (main text): 991, abstract 249; 1 figure; 2 tables**

22Potential competing interests

23Andrea Tricco, Dr Moher, and Raymond Daniel are members of the Cochrane
24Collaboration. Andrea Tricco is funded by a Canadian Institutes of Health Research
25Graduate Scholarship and a University of Ottawa Excellence Scholarship and Dr Moher
26is funded by a University of Ottawa Research Chair. These funders had no role in any of
27the following: study design; collection, analysis, and interpretation of data; writing of the
28report; and decision to submit the report for publication.

Background: Cochrane reviews are regarded as being scientifically rigorous, yet the proportion of Cochrane protocols that are published as reviews and the average time to publication remains unclear. If the time to publication of Cochrane reviews is affected by factors such as statistically significant results, this may lead to inaccurate decisions. We aimed to identify the frequency of published Cochrane reviews, determine their time to publication, and examine the factors associated with time to their publication.

Methods: Retrospective cohort study of Cochrane protocols published in Issue 2, 2000. The publication status of these reviews was tracked up to Issue 1, 2008 in The Cochrane Library. The time from protocol publication to the first review publication and review factors predicting the time to publication were analyzed via univariate and multivariate analyses.

Results: There were 130 new Cochrane protocols published in Issue 2, 2000. After excluding 12; 25/118 (21.2%) were unpublished and 93/118 (78.8%) were published as Cochrane reviews. Thirty-seven reviews (39.8%) were updates. The median time to publication was 1.6 years (range: 0.1-7.3 years). An author change between the protocol and final review was associated with longer time to publication ($p=0.002$), while an updated review was associated with shorter time to publication ($p=0.03$).

Conclusions: Only 80% of Cochrane protocols were published as final Cochrane reviews. The median time to publication was 19 months yet some reviews took much longer. Strategies to decrease time to publication should be considered, such as providing support to reviewers when a change in authorship occurs.

50Introduction

51Publication bias occurs when studies with certain characteristics (e.g., study funding)
52have a greater likelihood of being published (1), and being published quicker (2), than
53studies without these characteristics. Previously we examined publication bias of
54systematic reviews (SRs) through a retrospective cohort study and found that the majority
55of them (301/372; 81%) were published after 8 years of follow-up (3). A shorter time to
56publication was associated with the review being updated (hazard ratio: 1.80 [95%
57confidence interval: 1.39 to 2.33 years]) while a longer time to publication was associated
58with the review having two published protocols, indicating changes to the review plan
59(hazard ratio: 0.33 [95% confidence interval: 0.12 to 0.90 years]). We did not examine the
60*review factors* (e.g., statistically significant results) in our previous study. As such, we
61conducted this study to examine the association between review factors and the time to
62publication of Cochrane reviews.

63Methods

64All new protocols published in Issue 2, 2000 of the Cochrane Database of Systematic
65Reviews were selected. Their status was tracked through searching The Cochrane Library
66until January 23, 2008 (Issue 1, 2008) and contacting corresponding authors or the
67Cochrane Review Group coordinator. Cochrane protocols that were split into more than
68one Cochrane review, taken over by another review group, published in the same issue as
69the final Cochrane review, published later than the review publication or not published
70for the first time in Issue 2, 2000 were excluded (3).

71 A comprehensive data abstraction form was developed and pilot-tested. The form
72was used to abstract review characteristics (e.g., number of authors), methodology used

73by the reviewers (e.g., number of primary outcomes, inclusion of unpublished material,
74assessment of publication bias), and other factors (e.g., funding, number of updates).

75 In addition, the SR results and conclusions were classified using a system
76reported elsewhere (4). Briefly, results were classified as being non-statistically
77significant negative (e.g., unfavourable towards treatment intervention and $p>0.05$),
78statistically significant negative, neutral (i.e., effect size between 0.95 and 1.05 and the
79confidence interval crosses 1), non-statistically significant positive (e.g., favourable
80towards treatment intervention and $p>0.05$), statistically significant positive, and
81indeterminate (i.e., unable to judge; e.g., the SR lists 10 primary outcomes, all of which
82have different results). The conclusions were classified as being positive (i.e., authors
83stated that there is evidence of effectiveness), neutral (i.e., no evidence of effectiveness or
84they reported no opinion), negative (i.e., authors advised against use of the intervention
85or it was not recommended) or indeterminate (i.e., stated that there is insufficient
86evidence or that more research is required).

87 Published Cochrane reviews were analyzed via univariate and multivariate
88analyses. This was conducted by taking the log of the time between the “most recent
89substantive amendment date” of the protocol and subsequent review. Variables chosen for
90the univariate and multivariate analyses were based on *a priori* consideration. Statistical
91analyses were conducted with SAS, version 9.0 (SAS Institute, Cary, North Carolina).

92Results

93There were 130 new Cochrane protocols in Issue 2, 2000. After excluding 12; 25/118
94(21.2%) were unpublished and 93/118 (78.8%) were published as Cochrane reviews
95(Figure).

96 A primary outcome was reported in 80.6% (75/93) of the reviews, the majority
97 included all languages of publication (52.7%, 49/93), as well as published and
98 unpublished material (74.2%, 69/93; Table 1). Publication bias was found to affect the
99 results in only 11.1% (1/9) of the reviews. The majority of Cochrane reviews that
100 performed a meta-analysis of the primary outcome had statistically significant positive
101 results (31.5%, 17/54) while the majority of the conclusion statements were
102 indeterminate (43.0%, 40/93). Only 38.7% (36/93) of the reviews were subsequently
103 updated. A not-for-profit funding source was the most commonly reported funder (50.0%,
104 37/74).

105 The median time to publication was 1.60 years (range: 0.15 to 7.31 years). Four
106 factors significantly predicted publication (univariate analysis): the review having two
107 protocols, being subsequently updated, an author change between the protocol and
108 review, and number of included studies (all $p < 0.05$). An additional four factors were
109 included in the multivariate analyses due to *a priori* consideration: results, conclusions,
110 funding, and number of authors. Of the 8 factors, an author change between the protocol
111 and review was associated with longer time to publication ($p = 0.002$), and an updated
112 review was associated with shorter time to publication ($p = 0.03$; Table 2).

113 Discussion

114 Our results indicate that for every four published Cochrane reviews, one will remain
115 unpublished. Furthermore, a longer time to publication was associated with a change in
116 authorship between the Cochrane protocol and subsequent review. The Cochrane
117 Collaboration should consider strategies to improve the publication rate of Cochrane

118reviews as well as decrease their time to publication. These may include providing
119additional support to reviewers when an author change occurs.

120 Our study identified a large proportion of unpublished SRs. An international
121survey of systematic reviewers identified a 12.4% non-publication rate of SRs (5). In a
122retrospective cohort study of the protocol factors predicting publication of Cochrane
123reviews, only 13% (9/68) of the unpublished SRs were published elsewhere (3). This
124implies that a large amount of potentially meaningful data is missing from the literature
125and implies enormous wasted time and resources.

126 Our results are inconsistent with research examining publication bias of individual
127studies (e.g., trials) (2;6-10). In these studies, statistically significant results and funding
128were associated with publication of SRs. Since our study did not find such an association,
129it is apparent that differences exist in publication patterns between SRs and trials.

130 As reported elsewhere, our study has some limitations (3). Only one investigator
131abstracted all of the data, which could have led to inaccuracies. Assessing the primary
132outcome, results, and conclusions of the Cochrane reviews was often difficult and may
133have also led to error (4).

134 In conclusion, only about 80% of Cochrane protocols were published as complete
135Cochrane reviews within 19 months and some reviews took much longer to be published.
136Strategies to decrease time to publication should be considered, such as providing support
137to reviewers when a change in authorship occurs.

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139**Acknowledgements**

140This study was presented as a poster presentation at the 16th Cochrane Colloquium 2008
141in Freiberg, Germany. We thank Carmen Ng for her assistance with pilot-testing the data
142abstraction form, Dr. Sally Hopewell of the UK Cochrane Center for her expert advice on
143the study methods and for lending us the Cochrane CDs, as well as Lisa McGovern of the
144Canadian Cochrane Network and Centre for lending us the Cochrane CDs. ACT had full
145access to all of the data in the study and takes responsibility for the integrity of the data
146and the accuracy of the data analysis.

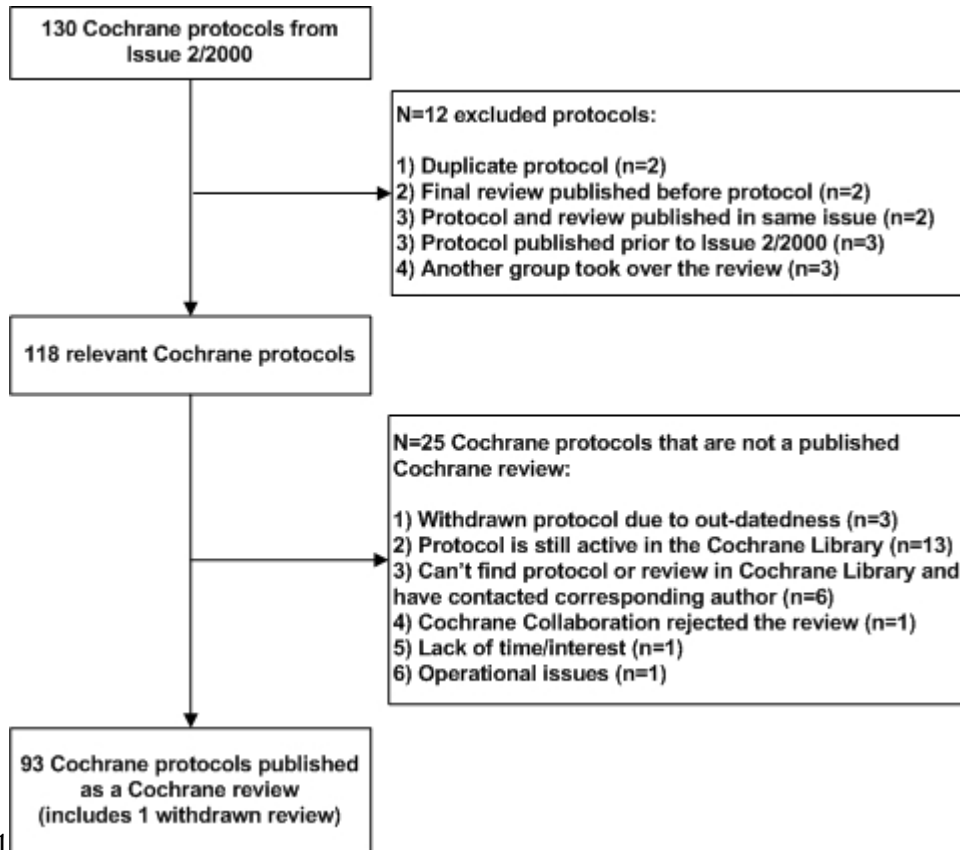
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148Author contributions:

149ACT conceptualized the research, obtained the sample of Cochrane reviews, designed the
150data abstraction form, abstracted all of the data from the reviews, verified the quality of
151the data, analyzed the results, wrote the manuscript, and approved the final version of the
152manuscript. ACT had full access to all the data in the study and takes full responsibility
153for the integrity of the data and the accuracy of the data analysis. DM conceptualized the
154research, designed the data abstraction form, edited the manuscript, and approved the
155final version of the manuscript. MHC verified the quality of the data, helped analyze the
156results, edited the manuscript, and approved the final version of the manuscript. RD
157helped obtain the sample of Cochrane reviews, edited the manuscript, and approved the
158final version of the manuscript.

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160Figure: Study Flow



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162Table 1: Cochrane review characteristics

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Item	Total: 93 published reviews
Descriptive characteristics	
Country of conduct: n (%)	
United Kingdom	36 (38.7)
Australia and New Zealand	20 (21.5)
Canada	7 (7.5)
United States of America	7 (7.5)
India	4 (4.3)
Italy	3 (3.1)
Netherlands	2 (2.2)
Denmark	2 (2.2)
Thailand	2 (2.2)
Other	10 (10.8)
Population examined: n (%)	
Neonates only	7 (7.5)
Children only	1 (1.1)
Adults only	20 (21.5)
Women only	13 (13.9)
Men only	1 (1.1)
Children and adolescents	1 (1.1)
Adolescents and adults	1 (1.1)
Adolescents, adults and elderly	1 (1.1)
Adults and elderly	1 (1.1)
All	47 (50.5)
Number of authors: median (range)	3 (1, 8)
Author change between protocol and corresponding review: n (%)	49 (52.7)
Review had two protocols: n (%)	2 (2.1)
Review had two “unique” Cochrane identification numbers: n, (%)	9 (9.7)
Number of studies included: median (range)	5 (0, 84)
Overall number of participants included: median (range)	410 (0, 109394)
Number of pages of pdf review file: median (range)	23 (5, 183)
Methodological characteristics	
Type of reports included in the review: n (%)	
Observational only	0 (0)
Experimental and quasi-experimental only	89 (95.7)
Both	4 (4.3)
Number of databases searched: median (range)	4 (1, 17)
A primary outcome was reported: n (%)	75 (80.6)
Number of primary outcomes: median (range)	1 (1, 20)
Languages included: n (%)	
English only	0 (0)
Mixed languages only	2 (2.1)
All languages	49 (52.7)
Not reported	42 (45.2)
Status of study report included: n (%)	
Published material only	2 (2.2)
Published and unpublished material	69 (74.2)
Not reported	22 (23.6)
Meta-analysis of the primary outcome was conducted: n (%)	54 (58.1)
Publication bias was assessed: n (%)*	9 (16.7)

Publication bias affected the results: n (%) [*]	1 (1.9)
Heterogeneity assessed: n (%) [*]	54 (100.0)
Heterogeneity affected the results: n (%) [*]	8 (14.8)
Results and conclusions classification	
Results [*]	
Indeterminate	9 (16.6)
Non-statistically significant negative	10 (18.5)
Statistically significant negative	0 (0)
Neutral	7 (13.0)
Non-statistically significant positive	11 (20.4)
Statistically significant positive	17 (31.5)
Conclusions	
Indeterminate	40 (43.0)
Negative	10 (10.8)
Neutral	24 (25.8)
Positive	19 (20.4)
Other factors	
Gender of corresponding author: n (%)	
Female	32 (34.4)
Male	48 (51.6)
Unclear	13 (14.0)
Corresponding author was a healthcare provider: n (%)	25 (26.9)
Number of reviews that were updated: n (%)	36 (38.7%)
Number of updates per review: median (range)	1 (1, 3)
Number of reviews with funding: n (%)	74 (80.0)
Type of funding source: n (%) [†]	
Government only	17 (23.0)
Not-for-profit organization only	37 (50.0)
Government and not-for-profit organization	19 (25.7)
For-profit and government and not-for-profit	1 (1.3)

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165**Notes:** * Denominator is number of reviews for which a meta-analysis was conducted (n=54), †

166denominator is number of reviews with funding (n=74).

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Table 2: Factors associated with the time to publication of Cochrane reviews from the multivariate analysis

Factor	Estimate*	Standard error	p-value
Two protocols (no vs. yes)	-0.810	0.467	0.086
Author change (no vs. yes)	-0.426	0.134	0.002 †
Number of authors	0.034	0.051	0.504
Funding (no vs. yes)	0.280	0.167	0.097
Number of included studies	0.007	0.005	0.145
Results (favourable vs. unfavourable)	0.018	0.274	0.948
Results (all others vs. unfavourable)	-0.167	0.244	0.495
Conclusions (negative vs. positive)	0.059	0.276	0.832
Conclusions (all others vs. positive)	-0.109	0.188	0.562
Update (no vs. yes)	0.305	0.138	0.030 †

Notes: * Negative value indicates longer time to publication and positive value indicates shorter time to publication, † p<0.05

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