

1 Searching clinical trial registries in interventional physical

2 therapy systematic reviews: A Pilot Cross-sectional Analysis

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- 10 **Abstract**
- 11 **Background.** Studies with positive findings are more likely to be published compared to those
- with negative findings. Therefore the latter studies are often disregarded in systematic reviews.
- 13 This causes an overestimation of a treatment effect size which leads to a misinterpretation of the
- evidence. Searching clinical trial registries in systematic reviews is a useful source to retrieve
- 15 unpublished clinical trials leading to the reduction of publication bias. Previous studies in the
- 16 literature reported inconsistent searching of clinical trial registries in systematic reviews
- published in several medical fields. Searching clinical trial registries in physical therapy is still
- unknown. The aim of this cross-sectional analysis is to evaluate the extent of clinical trial registry
- searching in physical therapy interventional systematic reviews.
- 20 **Methods.** Systematic reviews published between January 2017 and January 2018 were retrieved
- 21 from five reputable physical therapy journals. Interventional systematic reviews that were
- 22 coherent with the inclusion criteria were included in the analysis.
- 23 **Results.** The search yielded 40 systematic reviews. Among these 19 were interventional
- 24 systematic reviews as well as being consistent with the inclusion criteria and thus were
- considered for the analysis. After reviewing their search methodology, only two reviews (10.5%)
- 26 reported searching at least one clinical trial registry.
- 27 **Discussion.** The results of this study suggest poor searching of clinical trial registries in physical
- 28 therapy systematic reviews. Due to the limitations of this study, further research analyzing large
- 29 samples of interventional physical therapy systematic reviews is required.

Introduction

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- 3 Systematic reviews constitute a handy source of evidence for clinicians in the era of evidence
- 4 based practice(Gopalakrishnan & Ganeshkumar, 2013; Medina & Pailaquilén, 2010). This might
- 5 be due to their ability to summarize large numbers of trial findings (Mallett et al., 2012).
- 6 However, systematic reviews might be prone to publication bias (also known as selective
- 7 publishing of studies) represented in the consideration of published trials exhibiting significant
- 8 results with the disregard of unpublished studies exhibiting non-significant results(Onishi &
- 9 Furukawa, 2014). Thus this bias can be highly misleading to researchers and clinicians alike at
- interpreting the evidence(Gilbody et al., 2000; Joober et al., 2012) in the form of overestimation
- of the effects of a specific therapeutic intervention(Crawford, Briggs, & Engeland, 2010).
- 12 Therefore in order to prevent compromising the validity of systematic reviews, it is imperative
- for reviewers when conducting a systematic search to include a source that permits the retrieving
- of unpublished studies.
- 15 Searching clinical trial registries is one method utilized to retrieve unpublished studies which
- would help as well in the reduction of publication bias in systematic reviews. These registries
- promotes transparency in healthcare research via providing information about clinical trials being
- conducted irrespective of their results to the general public, patients, researchers, and clinicians.
- 19 Previous studies have shown that a search strategy targeting clinical trial registries is not
- 20 routinely present in systematic reviews in different medical fields (Combs, Atakpo, & Vassar,
- 21 2018 ;Jones et al., 2014 ;Keil, Platts-Mills, & Jones, 2015 ;Sinnett et al., 2015). In general
- medicine, a cross sectional study reported that 35% of systematic reviews published in this field
- stated the utilization of a search strategy targeting at least one clinical trial registry(Jones et al.,
- 24 2014). This percentage was even lower in systematic reviews published in the fields of
- emergency medicine (20%) (Keil et al., 2015), obstetrics and gynecology (18.4%) (Bibens,
- 26 Chong, & Vassar, 2016), dermatology (9.7%) (Combs et al., 2018), and clinical neurology
- 27 (6.3%) (Sinnett et al., 2015).
- 28 Concerning clinical trial registration in the physical therapy field, one study reported that only
- 29 about 34% of randomized controlled trials published in 2009 and indexed in the Physiotherapy
- 30 Evidence Database (PEDro) were registered (Pinto et al., 2013). In another study however clinical
- 31 trial registration of trials published in major physical therapy journals reported a growth from
- 4.3% in 2008 to 48.2% in 2012(Babu et al., 2014). This growth was attributed due to editorial
- policy recommendations set by multiple physical therapy journals (Babu et al., 2014).
- To the author's knowledge, there exists no published study in the literature that measured the
- 35 searching of clinical trial registries in physical therapy systematic reviews. Thus the aim of this
- 36 cross-sectional study is to examine the extent of searching clinical trial registries in therapeutic
- 37 systematic reviews published in selected reputable physical therapy journals. The author



- 1 hypothesize that searching clinical trial registries will be underutilized in physical therapy
- 2 systematic reviews in a similar fashion to systematic reviews in other medical fields.

3 **Method**

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4 Data Source

- 5 Interventional systematic reviews that were identified were published between January 2017 and
- 6 January 2018 in five selected reputable journals in the field of physical therapy. These journals
- 7 consisted of: Physical Therapy, Journal of Physiotherapy, Physiotherapy, Journal of Orthopedic
- 8 and Sports Physical Therapy, and Journal of Neurologic Physical Therapy. The reason for
- 9 selecting the previously mentioned journals is due to the popularity of these journals and thus
- 10 influencing greatly the physical therapy practice worldwide. Corrigendum of previously
- published systematic reviews before January 2017 will not be considered in the analysis.
- 12 The criteria of including systematic reviews in the analysis are systematic reviews having a
- predefined clinical question, an explicitly stated inclusion and exclusion criteria, a detailed
- description of the search strategy utilized, and finally having tested the effectiveness of a specific
- physical therapy treatment option on a certain condition or illness.

Study Selection Process

- 17 The author conducted a search of the journal issues published between January 2017 and January
- 18 2018 to retrieve systematic reviews. The manuscripts will then be screened and systematic
- 19 reviews that are compliant with the inclusion criteria will be accepted in the analysis. In case the
- 20 title of the manuscript did not explicitly include the words "systematic review" and/or "meta-
- analysis", the abstract of the manuscript will be reviewed in order to determine whether the
- 22 manuscript is a systematic review. After that, the full text of the review will be screened to
- 23 determine if the review is consistent with the criteria set previously. Therefore supplement issues
- 24 containing conference abstracts will be disregarded due to the absence of full texts. In case of
- 25 consistency, the review will be accepted in the analysis. Systematic reviews employing a search
- 26 strategy of databases mentioned in previously published reviews or protocols of reviews will as
- well be included in the analysis.

28 **Outcome Measures**

- 29 Following the inclusion of interventional systematic reviews, the author will reexamine the
- methods section not to mention the online supplements and appendices (if present) of each of the
- 31 included systematic reviews to determine if at least one clinical trial registry was included in the
- 32 systematic search. The clinical trial registries included as outcome measures were
- 33 ClinicalTrials.gov and 17 primary clinical trial registries in the World Health Organization
- 34 (WHO) Registry Network that meets the requirements of the International Committee of Medical
- 35 Journal Editors.
- 36 In case the authors of an included systematic review included a search strategy targeting at least
- one of the previously mentioned clinical trial registries, information regarding the name of the



- 1 registry, the number of studies retrieved from searching the registries will be extracted, not to
- 2 mention whether these studies were included or excluded by the review (and if the publication
- 3 status of the registered trials was the cause of exclusion).

Results

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- 5 Searching the previously mentioned five physical journals from January 2017 till January 2018
- 6 yielded 40 systematic reviews. After reviewing the full text of these studies, it was determined
- that only 19 were interventional systematic reviews and were thus included in the analysis.
- 8 After examining the search strategy of these interventional systematic reviews, it was revealed
- 9 that only two systematic review(Hall et al., 2017; Medeiros et al., 2017) of the included 19
- 10 (about 10.5%) had search strategies encompassing clinical trial registries. (figure 1)
- One (Medeiros et al., 2017) of these was published in the Journal of Physiotherapy and the
- other(Hall et al., 2017) was published in the *Physiotherapy* journal. The systematic review
- published by Medeiros and colleagues (Medeiros et al., 2017) searched Clinical Trials.gov while
- the other systematic review published by Hall and colleagues(Hall et al., 2017) searched the
- 15 International Standard Randomised Controlled Trial Number (ISRCTN).
- None of the previously mentioned systematic reviews did declare information about the number
- of clinical trials that were retrieved by this search. Thus no additional information can be
- 18 extracted whether these retrieved studies were included or excluded on the basis of their
- 19 publication status.

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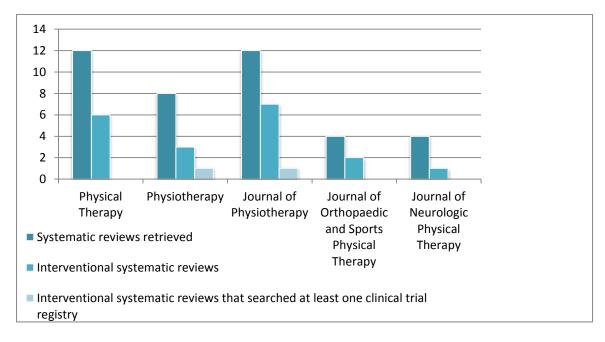


Figure 1 shows number of systematic reviews retrieved, the number of interventional systematic reviews and the reviews that searched at least one clinical trial registry



Discussion

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- 2 Since systematic reviews constitute have a major role of influencing the practice of physical
- 3 therapy clinicians worldwide, authors of these reviews should make use of all trial findings
- 4 regardless of their publication status in order to minimize the harmful effect of publication bias.
- 5 One the various methods of reducing publication bias in systematic reviews is to search for
- 6 unpublished studies via clinical trial registries. This study is a novel cross sectional analysis of
- 7 physical therapy systematic reviews published between January 2017 and January 2018. Five
- 8 reputable physical therapy journals were selected as a source to retrieve eligible interventional
- 9 systematic reviews from. Although no similar study has been performed in the physical therapy
- field, multiple studies have examined the extent of searching clinical trial registries in systematic
- 11 reviews in other health related fields.
- 12 The results show that clinical trial registries are poorly searched in physical therapy systematic
- reviews. However this study is not prone to limitations. The most important limitation in this
- cross-sectional study is the small number of physical therapy journals being examined. There
- exist more physical therapy journals that were not included in the study. In addition to that,
- multidisciplinary rehabilitation and sports medicine journals were not examined. Therefore it
- 17 might be difficult to generalize the results obtained to all published physical therapy
- interventional systematic reviews. Another limitation of the study is characterized in the absence
- of more than one author to retrieve, include, and examine the systematic reviews. This cross-
- sectional study only focused on systematic reviews published between January 2017 and January
- 21 2018 in the English language. Thus, the results obtained cannot be generalized to physical
- therapy interventional systematic reviews published before January 2017 and after January 2018
- and to reviews published in non-English languages.

24 **Conclusion**

- 25 Clinical trial registries were not consistently utilized in interventional physical therapy systematic
- 26 reviews published between January 2017 and January 2018. Due to the vast limitations of the
- study, future research should address the limitations listed previously and therefore employ a
- 28 strategy that shall retrieve larger numbers of physical therapy interventional systematic reviews
- 29 to be examined. In addition to that, reviewers conducting systematic searches in the field of
- 30 physical therapy should be encouraged to include a strategy that permits them to search clinical
- 31 trial registries.

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