

Quality of pharmaco-economic research in China

A systematic review

Huifen Ma, PhD^a, Weiyan Jian, PhD^b, Tingting Xu, MS^a, Yasheng He, MS^a, John A. Rizzo, PhD^c, Hai Fang, PhD^{a,*}

Abstract

Background: The number of pharmaco-economic publications in the literature from China has risen rapidly, but the quality of pharmaco-economic publications from China has not been analyzed.

Objectives: This study aims to identify all recent pharmaco-economic publications from China, to critically appraise the reporting quality, and to summarize the results.

Methods: Four databases (PubMed, Web of Science, Medline, and EmBase) were searched for original articles published up to December 31, 2014. The Consolidated Health Economic Evaluation Reporting Standards statement including 24 items was used to assess the quality of reporting of these articles.

Results: Of 1046 articles identified, 32 studies fulfilled the inclusion criteria. They were published in 23 different journals. Quality of reporting varied between studies, with an average score of 18.7 (SD=4.33) out of 24 (range 9–23.5). There was an increasing trend of pharmaco-economic publications and reporting quality over years from 2003 to 2014. According to the Consolidated Health Economic Evaluation Reporting Standards, the reporting quality for the items including “title,” “comparators of method,” and “measurement of effectiveness” are quite low, with less than 50% of studies fully satisfying these reporting standards. In contrast, reporting was good for the items including “introduction,” “study perspective,” “choice of health outcomes,” “study parameters,” “characterizing heterogeneity,” and “discussion,” with more than 75% of the articles satisfying these reporting criteria. The remaining items fell in between these 2 extremes, with 50% to 75% of studies satisfying these criteria.

Conclusion: Our study suggests the need for improvement in a number of reporting criteria. But the criteria for which reporting quality was low seem to be limitations that would be straightforward to correct in future studies.

Abbreviations: CHEERS = Consolidated Health Economic Evaluation Reporting Standards, ISPOR = The International Society for Pharmacoeconomics and Outcomes Research, QHES = Quality of Health Economic Studies.

Keywords: China, pharmaco-economic, research quality

Editor: Bernhard Schaller.

WJ is the co-first author with equal contributions.

Author contributions: HF, HM, and JR planned the study. HM, WJ, TX, and YH conducted the search, extracted, analyzed and interpreted the data, and produced a draft of the manuscript. HF and JR oversaw the progression of the review, provided guidance and contributed to various versions of the manuscript. All authors read and approved the final manuscript.

Funding disclosures: Financial support for this study was provided to Dr Hai Fang by a grant from National Natural Science Foundation of China (Grant Number 71373013) and a grant from Peking University Health Science Center (Grant Number BJMU20130338). The funding agreement ensured the authors' independence in designing the study, interpreting the data, writing, and publishing the article.

None of the authors have expressed any conflict of interest.

^a China Center for Health Development Studies, Peking University, ^b Department of Health Policy and Administration, Peking University, Haidian District, Beijing, China, ^c Departments of Economics and Department of Preventive Medicine, State University of New York at Stony Brook, Stony Brook, NY.

* Correspondence: Hai Fang, China Center for Health Development Studies, Peking University, Beijing, China (e-mail: hfang@hsc.pku.edu.cn).

Copyright © 2016 the Author(s). Published by Wolters Kluwer Health, Inc. All rights reserved.

This is an open access article distributed under the terms of the Creative Commons Attribution-Non Commercial-No Derivatives License 4.0 (CCBY-NC-ND), where it is permissible to download and share the work provided it is properly cited. The work cannot be changed in any way or used commercially.

Medicine (2016) 95:41(e5114)

Received: 10 May 2016 / Received in final form: 12 September 2016 /

Accepted: 18 September 2016

<http://dx.doi.org/10.1097/MD.0000000000005114>

1. Introduction

Pharmacoeconomics refers to the scientific discipline that compares the value of 1 pharmaceutical drug or drug therapy to another.^[1,2] The number of pharmaco-economic publications in the literature from China has risen rapidly.^[3] Ideally, these studies should follow evaluation criteria commonly accepted by researchers in this field.^[4,5] This is critical for helping to ensure the quality and reliability of the research. Several studies have examined trends in pharmaco-economic publications from China in terms of publication numbers, journal placement, and the authors and universities involved in the research.^[6] To our knowledge, however, no study has yet analyzed the “quality” of pharmaco-economic publications from China.

Evaluating the quality of pharmaco-economic research is challenging. These studies are complex, often including several treatments or interventions, and may involve detailed clinical pathways. Various cost and effectiveness measures must be obtained as well. A variety of issues, including the perspective of the study, economic modeling and assumptions, addressing uncertainty, and so on, present significant challenges to editors and reviewers in judging the quality of a pharmaco-economic study. As a result, it is perhaps not surprising that the quality of pharmaco-economic publications in the literature varies substantially.^[7]

Despite its complexity, pharmaco-economic research offers important information for health care and health policy decision makers to help allocate healthcare resources more efficiently.

Until relatively recently, there was a paucity of pharmaco-economic research to guide policy making in China, but the number of such studies has risen rapidly. China has implemented major healthcare system reforms since 2009, and pharmaco-economic studies have become increasingly critical in light of these changes to inform policies aimed at controlling costs while enhancing quality of care. Thus, it is important to assess the quality of this growing literature in pharmaco-economic research. The present study seeks to perform such an evaluation, to identify any weaknesses in existing research and to provide recommendations for ensuring the quality and integrity of future pharmaco-economic research in China.

The present study will evaluate the quality of pharmaco-economic papers published in English, but conducted in China. Whereas there are more pharmaco-economic publications in Chinese than in English, English journals are on average more impactful. Moreover, English publications are becoming an increasingly important outlet for pharmaco-economic research in China. Vaccines are excluded from the present study, because they have not been regarded as pharmaceutical products in China.

Guidelines for performing appropriate pharmaco-economic studies have been well delineated. The present study employs a recently developed health economic quality criterion: Consolidated Health Economic Evaluation Reporting Standards (CHEERS) statement to evaluate the quality of pharmaco-economic studies examined in this review.^[8]

2. Methods

2.1. Literature search

A systematic search of the literature was conducted in March 2015 using PubMed, Web of Science, Medline, and EmBase to identify pharmaco-economic studies pertaining to China. We found a total of 1046 potential articles by this initial search. Search terms in all 4 databases included “Drug economic,” “pharmaco-economic,” “Economics, Pharmaceutical,” “economics,” “pharmacy,” “pharmaceuticals,” “health economic,” “Medical Economics,” “cost,” “Cost Measures,” “cost-effectiveness analysis,” “cost-minimization analysis,” “cost-utility analysis,” “cost-benefit analysis,” “Benefits and Costs,” “Data,” “Cost-Benefit,” and “China.” These key words were used alone and in indifferent combinations. The exclusion criteria were as follows: duplicated articles; not pharmaco-economic studies; studies comparing multiple countries, not only about China; not full-journal articles, such as meeting abstracts, letters to the editor, treatment guidelines or recommendations, expert opinion, and narrative reviews. Studies comparing multiple countries were also excluded.

Two researchers carried out the literature search using the English-based search engines and identified articles independently. They assessed the abstracts of the identified studies, and all abstracts that met the inclusion criteria were confirmed by a third researcher. Full articles were then obtained for further evaluation.

We used NoteExpress V3.0 to review and evaluate the included studies. First, all the searched articles were compiled into NoteExpress. Second, 2 reviewers simultaneously screened the articles by titles and abstracts according to the exclusion criteria. Third, the full texts of the included articles were downloaded into NoteExpress. Finally, the full texts of the included articles were reviewed by 3 researchers.

Ethical approval was not necessary, because the present systematic review did not involve patients. The present article reviewed the previous publications about quality of pharmaco-

economic research in China, and all the materials were from the previous publications.

2.2. Evaluation of studies

There are a number of published criteria for evaluating pharmaco-economic research. In the 1970s, Alan Williams at the University of York developed the first widely adopted health economic evaluation guidelines.^[9] In 1987, Michael Drummond suggested 10 health economic standards, which have been widely accepted by researchers in the field of health economics. In 1995, the *British Medical Journal* developed guidelines for pharmaco-economic studies.^[10] Each of these evaluation standards or guidelines are extremely helpful for health economic research and their quality improvement. In 2003, a Quality of Health Economic Studies (QHES) instrument was designed to evaluate all 3 common types of health economic analyses: cost-minimization, cost-effectiveness, and cost-utility.^[11,12] The instrument emphasizes appropriate methods, valid and transparent results, and comprehensive reporting of results in each study. The present study uses “The CHEERS.

The International Society for Pharmacoeconomics and Outcomes Research (ISPOR) introduced CHEERS statement, and it has been endorsed and published by the 10 publications.^[13] The ISPOR Health Economic Evaluation Publication Guidelines Good Reporting Practices Task Force was approved by the ISPOR Board of Directors in 2009, and it aimed to develop guidance to improve the reporting of health economic evaluations. Task force membership was comprised of health economic journal editors and content experts from around the world. Forty-seven participants representing academic, biomedical journal editors, the pharmaceutical industry, government decision makers, and those in clinical practice were invited to the 2-round Delphi Panel. The task force submitted their first draft to the ISPOR Health Economic Evaluation Publication Guidelines Good Reporting Practices Task Force Review Group, and 24 reviewers submitted written comments. The report was revised and re-titled CHEERS in May 2012, and the revised CHEERS report was presented at the ISPOR 17th Annual International Meeting in Washington, DC.^[14]

The CHEERS statement attempts to consolidate and update previous efforts into a single useful reporting standard. The CHEERS statement is not intended to prescribe how economic evaluations should be conducted; rather, analysts should have the freedom to innovate or make their own methodological choices. Its objective is to ensure that these choices are clear to reviewers and readers. The present study uses CHEERS to evaluate the quality of pharmaco-economic publications from China as it is the most recent and comprehensive guideline for this purpose.

The CHEERS is a 24-item scale covering 6 main categories: title and abstract, introduction, methods, results, discussion, and others. To estimate a summary reporting score, it is suggested to assign a value of 1 if the study fulfilled the requirement of reporting for that item completely, 0.5 for partially completing the requirement, and otherwise 0. Therefore, the maximum score for a publication that reports completely according to these standards is 24. At least 2 reviewers independently appraised the studies considered in this review. When results differed, they were discussed by all 4 researchers until any discrepancies were resolved.

2.3. Statistical analysis

In presenting our findings, we report the number of publications in each year, the country of the first author, and the publication

journals. As noted above, the CHEERS is used to assess the quality of pharmaco-economic publications from China up to December 2014.

3. Results

3.1. Study selection process

Figure 1 shows the flowchart for searching pharmaco-economic publications from China. The initial database search identified 1046 articles. After screening by title and abstract, 178 full economic evaluations were identified. Of those, only 32 satisfied study inclusion criteria. Studies were mainly excluded because they were duplicated results (the same articles searched from different databases, n=386), not pharmaco-economic studies (n=532), not about China (n=39), or were themselves review articles (n=57). It is quite common to make the initial search very broad to avoid the possibility of omitting any relevant studies, but the final number of studies satisfying inclusion criteria is typically much smaller than the initial search identified.^[6,15-16]

3.2. Overview of included studies

A description of the 32 articles is presented in Table 1. These studies were published between 2003 and 2014. There have been more publications in recent years. In 2013, there were 8 publications, and in 2012 and 2014, there were 5 publications in each year. The first authors are mainly from China (28 of

32 publications, 87.5%). These studies appeared in a variety of journals. Nineteen (59.3%) papers were published in journals from the United States. The majority of publications were in medical journals (25 publications or 78.1%). The rest appeared in health economic journals or health services research journals (7 publications or 21.9%).

3.3. Results of the quality of reporting assessment

The results of the assessment of reporting quality per study are summarized in Table 2 and Table 3. The complete references of these 32 articles were reported in the reference list of 26 to 57. Substantial differences in the quality of reporting were observed among articles with an average score of 18.7 (SD=4.33) out of 24 (range 9–23.5). Figure 2 shows the average score for

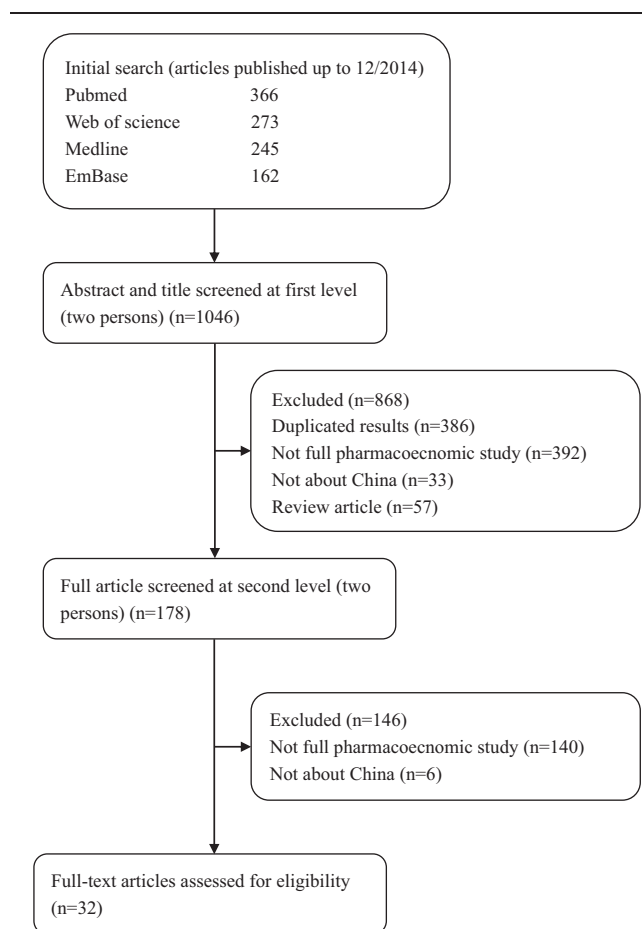


Figure 1. Flowchart of search results and selection process.

Table 1
Description of articles (n=32).

Categories	n (%)
Published year	
2003	2 (6.3)
2006	1 (3.1)
2008	3 (9.4)
2009	4 (12.5)
2010	2 (6.3)
2011	2 (6.3)
2012	5 (15.6)
2013	8 (25.0)
2014	5 (15.6)
Country of residence of the first author	
China	28 (87.5)
Other	4 (12.5)
Journal	
Value in Health	6 (18.8)
PloS One	4 (12.5)
Clinical Therapeutics	3 (9.4)
Advances in Therapy	1 (3.1)
Alimentary Pharmacology & Therapeutics	1 (3.1)
Asia-Pacific Journal of Clinical Oncology	1 (3.1)
BMC Cancer	1 (3.1)
BMC Health Services Research	1 (3.1)
Cardiovascular Drugs and Therapy	1 (3.1)
Chemotherapy	1 (3.1)
Circulation. Cardiovascular Quality and Outcomes	1 (3.1)
Gastrointestinal Endoscopy	1 (3.1)
International Journal of Clinical Pharmacology and Therapeutics	1 (3.1)
International Journal of Hematology	1 (3.1)
International Journal of Technology Assessment in Health Care	1 (3.1)
International Urology and Nephrology	1 (3.1)
Journal of Digestive Diseases	1 (3.1)
Journal of General Internal Medicine	1 (3.1)
Medicina (Kaunas, Lithuania)	1 (3.1)
PharmacoEconomics	1 (3.1)
Radiology	1 (3.1)
The Australian and New Zealand Journal of Obstetrics and Gynaecology	1 (3.1)
Country of journal	
United States	19 (59.3)
United Kingdom	4 (12.5)
Australia	3 (9.4)
Germany	1 (3.1)
Japan	1 (3.1)
Lithuania	1 (3.1)
Netherlands	1 (3.1)
New Zealand	1 (3.1)
Switzerland	1 (3.1)

Table 2
Quality of pharmacoeconomic publications (articles 1–16).

CHEERS item	CHEERS item no.	Article number in the reference list															
		[17]	[18]	[19]	[20]	[21]	[22]	[23]	[24]	[25]	[26]	[27]	[28]	[29]	[30]	[31]	[32]
Title and abstract																	
Title	1	Yes	Yes	Part	Yes	Yes	Part	Part	Yes	Yes	Part	Yes	Yes	Yes	Yes	Yes	Yes
Abstract	2	Part	Yes	Part	Part	Part	Part	Yes	Yes	Yes	Yes	Yes	Yes	Part	Yes	Part	Part
Introduction																	
Background and objectives	3	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Part	Yes	Yes	Yes	Yes	Yes
Methods																	
Target population and subgroups	4	Yes	Part	Yes	Yes	Yes	Yes	Part	Part	Part	Yes	Part	Part	Yes	No	Yes	Part
Setting and location	5	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Part	Yes	Yes	Yes	Yes	No	Yes	Part
Study perspective	6	No	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Comparators	7	Part	Part	Yes	Yes	Part	Yes	Part	Part	Yes	Yes	Part	Part	Part	No	Yes	Part
Time horizon	8	Part	Yes	Yes	Yes	Yes	Yes	Yes	Part	No	Yes	Yes	Yes	Yes	Part	Yes	Part
Discount rate	9	No	Yes	Yes	No	Yes	No	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Choice of health outcomes	10	Part	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Part	Yes	Yes	Yes	Yes
Measurement of effectiveness	11a	NA	NA	NA	NA	NA	Yes	Part	NA	NA	NA	NA	NA	NA	NA	NA	NA
	11b	No	Yes	Yes	Yes	Part	NA	NA	Yes	Yes	Part	Yes	Part	Part	Yes	Yes	Part
Measurement and valuation of preference-based outcomes	12	No	Yes	Yes	Yes	Yes	Part	Yes	Yes	No	Yes	Yes	Yes	Yes	No	Yes	Yes
Estimating resources and costs	13a	No	NA	NA	Part	NA	Part	NA	NA	Part	NA	NA	NA	NA	NA	NA	Yes
	13b	NA	Yes	Yes	NA	Yes	NA	Yes	Yes	NA	Yes	Yes	Yes	Yes	Yes	Yes	NA
Currency, price date, and conversion	14	No	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Part	Yes	Yes	Yes	No	Yes	No
Choice of model	15	No	Yes	Yes	No	Yes	Part	Yes	Yes	No	Yes	Yes	Part	Yes	Yes	No	No
Assumptions	16	No	Yes	Yes	Yes	No	Part	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes	No	No
Analytic methods	17	Part	Yes	Yes	Part	Yes	Yes	Yes	Yes	Part	Yes	Yes	Yes	Yes	Yes	Yes	Part
Results																	
Study parameters	18	Part	Yes	Yes	Yes	Part	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Part
Incremental costs and outcomes	19	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes	Part	No	Yes
Characterizing uncertainty	20a	Part	NA	NA	Part	NA	Yes	NA	NA	Part	NA	NA	NA	NA	NA	NA	Part
	20b	NA	Yes	Yes	NA	Yes	NA	Yes	Yes	NA	Yes	Yes	Yes	Yes	Part	Yes	NA
Characterizing heterogeneity	21	Part	Yes	Yes	Part	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Part
Discussion																	
Study findings, limitations, generalizability, and current knowledge	22	Part	Yes	Yes	Part	Yes	Yes	Yes	Yes	Part	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Other																	
Source of funding	23	No	Yes	Yes	Yes	No	No	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes
Conflicts of interest	24	No	Yes	Yes	Yes	Yes	No	No	Yes	No	Yes	No	Yes	Yes	Yes	Yes	Yes
Scores		9.5	23	22	18	20	17	21	22.5	14	22.5	20.5	21.5	22.5	17.5	20.5	16

CHEERS = Consolidated Health Economic Evaluation Reporting Standards, NA = not applicable, no = not reported, part = partially reported, yes = reported.

pharmacoeconomic publications by year. Figure 3 shows each item of 24 reporting assessments by 3 categories: completely adequate, partially adequate, or inadequate.

Several observations are worth noting in above results. The reporting quality for item 1, and item 7 and item 11 are quite low—less than 50% of studies fully satisfied these criteria. We discuss each of these items in turn.

Item 1 is to identify the study as an economic evaluation, or use more specific terms such as “cost-effectiveness analysis” and describe the interventions compared. Only 15 publications specifically identify their study as economic evaluations and the drugs compared in their studies. Another 15 publications are partially adequate, and 2 publications do not report the type of economic evaluation. In addition, the titles of some articles may not correctly describe the types of studies.

Item 7 is to describe the interventions or strategies being compared and state why they were chosen. Thirteen publications fully satisfied this item, 18 publications partially satisfied it, and 1 study failed to satisfy it. Most publications that did not fully satisfy this item did not specify why the alternative drugs should be compared in the studies.

Item 11a is to describe fully the design features of the single effectiveness study and why the single study was a sufficient source of clinical effectiveness data. There are only 5 publications relevant to this item: 2 publications fully satisfied this criterion and 3 partially satisfied it.

Item 11b is to describe fully the methods used for identification of included studies and syntheses of clinical effectiveness data. There are 27 publications relevant to this item. Twelve publications fully met this criterion, 13 publications partially satisfied it, and 2 publications failed to satisfy it.

In contrast, reporting was relatively complete for items 3 and item 22, with at least 28 articles satisfying these criteria. More than 75% publications fully satisfied the following items: item 3, item 6, item 10, item 18, item 21, and item 22.

Item 3 is to provide explicit statements of the broader context for the study. Thirty publications not only satisfied this criterion but also presented the study question and its relevance for health policy or practice decisions. Only 2 publications were scored as partially satisfying this criterion because they did not present the relevance of the study for health policy or practice decisions.

Table 3
Quality of pharmacoeconomic publications (articles 17–32).

CHEERS Item	CHEERS item no.	Article number in the reference list															
		[33]	[34]	[35]	[36]	[37]	[38]	[39]	[40]	[41]	[42]	[43]	[44]	[45]	[46]	[47]	[48]
Title and abstract																	
Title	1	Yes	Part	Part	Part	Part	Part	Part	Part	Part	No	Part	Part	Yes	No	Part	Yes
Abstract	2	Part	Yes	Part	Yes	Yes	Yes	Part	Yes	Yes	Part	Part	Yes	Part	Part	Part	Yes
Introduction																	
Background and objectives	3	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Part	Yes	Yes	Yes	Yes	Yes	Yes
Methods																	
Target population and subgroups	4	Yes	Part	Yes	Yes	Part	Part	Yes	Yes	Yes	Part	Yes	Part	Yes	Yes	Part	Yes
Setting and location	5	Yes	Part	Yes	Part	No	Part	Yes	Part	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes
Study perspective	6	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	Yes	No	Yes	Yes	No	No	Yes
Comparators	7	Part	Part	Yes	Yes	Part	Part	Yes	Yes	Part	Part	Part	Yes	Part	Yes	Part	Yes
Time horizon	8	Part	Yes	Yes	Yes	No	Yes	Yes	Yes	No	Yes	Part	Part	No	No	Part	Yes
Discount rate	9	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes	No	No	No	No	Yes	No	No	Yes
Choice of health outcomes	10	Yes	Yes	Yes	Part	Yes	Yes	Part	Yes	Part	No	Yes	Yes	Yes	Part	Yes	Yes
Measurement of effectiveness	11a	NA	NA	NA	NA	NA	NA	NA	NA	Part	NA	Yes	NA	NA	Part	NA	NA
	11b	Yes	Part	Yes	Part	Part	Part	Part	Yes	NA	No	NA	Part	Part	NA	Part	Yes
Measurement and valuation of preference-based outcomes	12	Yes	Yes	Part	Yes	Yes	Yes	Part	Yes	No	No	No	Yes	Yes	Part	No	Yes
Estimating resources and costs	13a	NA	NA	NA	NA	NA	NA	NA	NA	Part	NA	Part	NA	NA	Part	NA	NA
	13b	Yes	Yes	Yes	Yes	Yes	Yes	Part	Yes	NA	Part	NA	Yes	Yes	NA	No	Yes
Currency, price date, and conversion	14	Yes	Part	Yes	Yes	Part	Yes	Yes	Yes	No	Part	No	Yes	Part	No	No	Yes
Choice of model	15	Part	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	No	No	Yes	Yes	No	Part	Yes
Assumptions	16	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	No	No	Yes	Yes	No	Part	Yes
Analytic methods	17	Yes	Yes	Part	Yes	Yes	Yes	Yes	Yes	Yes	Part	Yes	Yes	Yes	Part	Part	Yes
Results																	
Study parameters	18	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Part	Part	Yes	Yes	Yes	Part	Yes	Yes
Incremental costs and outcomes	19	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	No	Yes	Yes	Yes	Part	No	Yes
Characterizing uncertainty	20a	NA	NA	NA	NA	NA	NA	NA	NA	Part	NA	Yes	NA	NA	Part	NA	NA
	20b	Yes	Yes	Part	Yes	Yes	Yes	Yes	Yes	NA	No	NA	No	Yes	NA	Part	Yes
Characterizing heterogeneity	21	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Part	No	Yes	Yes	Yes	Part	Yes	Yes
Discussion																	
Study findings, limitations, generalizability, and current knowledge	22	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Part	Yes	Yes
Other																	
Source of funding	23	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	No	Yes	Yes	No	No	Yes
Conflicts of interest	24	No	Yes	Yes	Yes	Yes	No	Yes	Yes	No	Yes	Yes	No	Yes	No	Yes	Yes
Scores		21	21	20.5	22	19.5	20.5	21	23	11	9	14.5	19	20	9.5	12	23.5

CHEERS=Consolidated Health Economic Evaluation Reporting Standards, NA=not applicable, no=not reported, part=partially reported, yes=reported.

In item 22, 28 out of 32 publications summarized key study findings and described how they support the conclusions reached. They discussed limitations and the generalizability of the findings and how the findings fit with current knowledge. Four publications did not discuss generalizability of the findings or how the findings fit with current knowledge.

In item 6: 26 out of 32 publications described the perspective of the study and relate this to the costs being evaluated.

In item 10, 25 publications described what outcomes were used as the measures of benefit in the economic evaluation and their relevance for the type of analysis performed. Six were scored as partially satisfying this criterion because they did not present their relevance for the type of analysis performed. Only 1 publication completely failed to satisfy this criterion.

In item 18, the study parameters were clearly made by 26 out of 32 publications.

In item 21, 26 publications described characterizing heterogeneity and 1 publication did not describe it. Five publications did not report fully about the differences in costs, outcomes, or cost-effectiveness that can be explained by variations between subgroups of patients with different baseline characteristics.

The remaining items fell in between these 2 extremes, with 50% to 75% of studies satisfying these criteria.

4. Discussion

The assessment criteria used in the present study—CHEERS—have been applied in a number of systematic reviews reporting the quality of pharmacoeconomic publications.^[49–51] Our findings confirm that there have been an increasing number of pharmacoeconomic publications from China during the period of 2003 to 2014, and the reporting quality has also been improving significantly over time. Pharmacoeconomics has received greater attention in China, which is particularly important for resource allocations and health policy decision making. This study reviews and summarizes the current state of pharmacoeconomic research and provides insight into how to improve the quality of such research in China. The results of our review may provide health policymakers with a sense of the reliability of existing research to help inform their decisions. Our study suggests that the potential for improving the quality of pharmacoeconomic analyses to inform health policy is substantial.

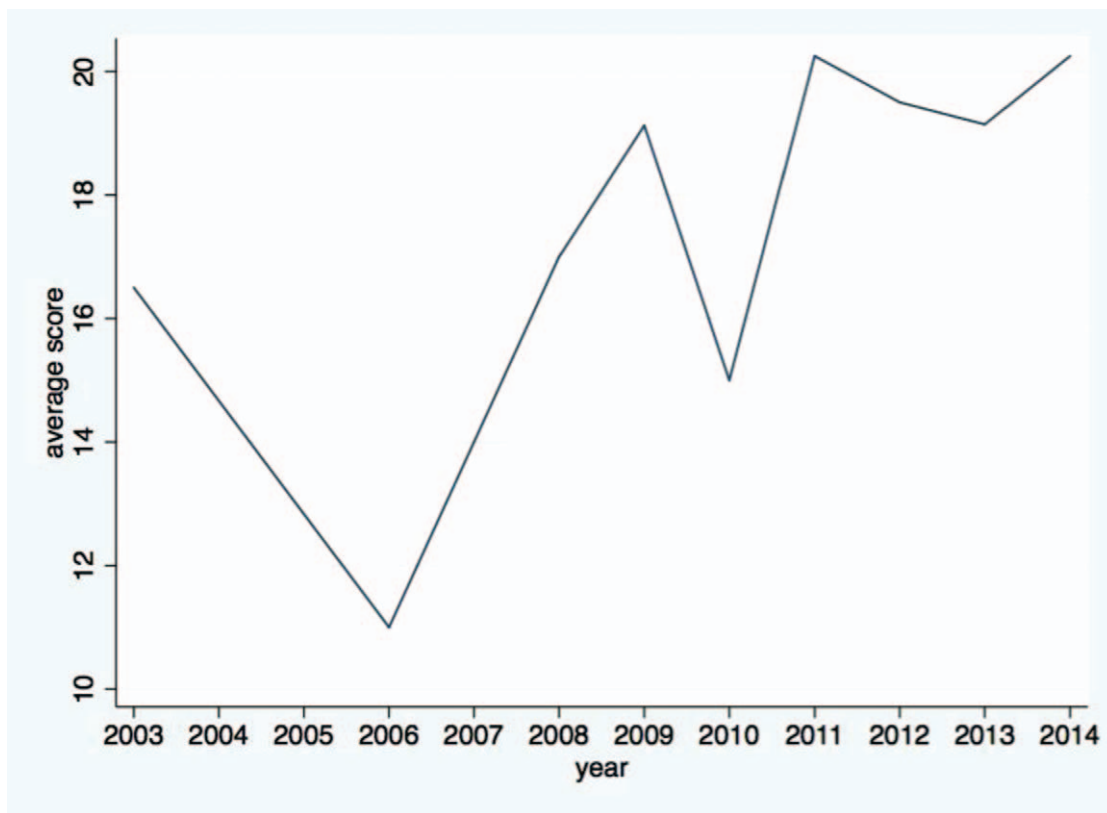
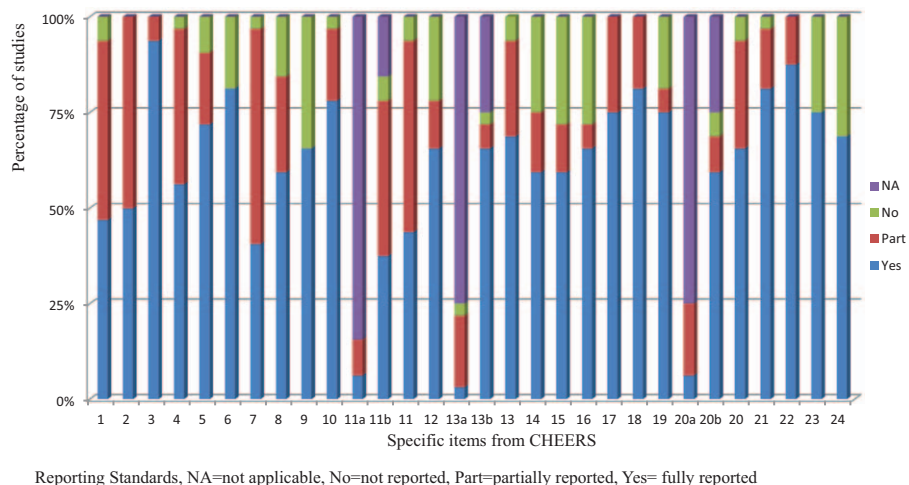


Figure 2. Average score of the publications by year.

Some systematic reviews have studied the quality of pharmacoeconomic publications from other countries.^[52-54] Desai et al^[20] examined the quality of pharmacoeconomic studies in India with 29 articles, and found that the lack of sensitivity analyses and discounting was the most important problem. Woersching et al^[21] assessed the quality of economic evaluations of US Food and Drug Evaluation (FDA) novel drug approvals as a systematic review including 36 articles and found that a minority

of the 2008 to 2009 novel drugs had mixed study quality. Gavaza et al^[54] studied the state of health economic (including pharmacoeconomic) research with 44 articles in Nigeria and concluded that the conduct of health economic (including pharmacoeconomic) research in Nigeria was limited, and about two-thirds of published articles were of suboptimal quality. The number of articles included in the above systematic reviews is in the range of 20 to 40, and the present study reviews 32 articles.



Reporting Standards, NA=not applicable, No=not reported, Part=partially reported, Yes= fully reported

Figure 3. Quality of publications per items of the CHEERS checklist. CHEERS=Consolidated Health Economic Evaluation Reporting Standards.

4.1. Quality of the studies

A lack of clarity in some basic items makes it difficult for readers to understand study objectives and to judge the quality and appropriateness of results. Although most published studies from China in our review have adhered to CHEERS, there are still a significant number of publications with low reporting quality. Significant items that affected the quality of economic studies included “Title,” “Comparators,” and “Measurement of effectiveness.” Our study suggests the need for improvement in these areas. Fortunately, these would seem to be reporting limitations that would be straightforward to correct in future studies.

Chinese authors should try to identify the study as an economic evaluation or use more specific terms such as “cost-effectiveness analysis,” and describe why they chose the compared interventions. In addition, some Chinese authors did not provide adequate information for the measurement of effectiveness.

There are approximately 1 million publications each year in the world.^[55] If the article titles are clear and indicate the nature of health economic studies, it will be much easier for researchers to search for them electronically and to compare results across similar studies. Previous studies have shown that a structured abstract is preferable to a descriptive structure for researchers to understand the main contents in a publication.^[56,57] In comparative studies, the choices of alternative interventions or drugs are particularly important to the results and conclusions of a study. The authors also need to explain the criteria for choosing the alternatives drugs that are being compared. Pharmacoeconomic publications from China have often omitted this important item, which is a serious shortcoming at present.

4.2. Limitations

The present study has several limitations that must be noted. First, the researchers are not blind to the authors and journals of pharmacoeconomic publications, so the assessment of reporting quality may be related to the personal opinions of reviewers. Second, all 24 items are treated equally, but some items may be more important in evaluating quality. Finally, the present study does not assess the scientific quality of pharmacoeconomic publications, as it only studies the reporting quality instead. Although reporting quality and scientific quality may be related, they are not the same thing.

5. Conclusions

Our systematic review identified 32 economic evaluations of drugs published between 2003 and 2014. Clarity of reporting should be an important part of every pharmacoeconomic study. Medical journals, particularly those with limited experience publishing pharmacoeconomic analyses, should adopt and enforce standard protocols for conducting and reporting this research.

References

- Mueller C, Schur C, O’Connell J. Prescription drug spending: the impact of age and chronic disease status. *Am J Public Health* 1997;87:1626–9.
- Brown GC, Brown MM. Value-based medicine and pharmacoeconomics. *Dev Ophthalmol* 2016;55:381–90.
- Rodrigues J, Wu JH, Clay E, et al. Impact of pharmacoeconomics guidelines on the international publications in China. *Value Health* 2014;17:A799.
- Drummond MF, Sculpher MJ, Torrance GW, et al. *Methods for the Economic Evaluation of Health Care Programmes*. 3rd ed. New York: Oxford University Press; 2005.
- Husereau D, Drummond M, Petrou S, et al. Consolidated Health Economic Evaluation Reporting Standards (CHEERS) statement. *Int J Technol Assess Health care* 2013;29:117–22.
- Jiang S, Ma X, Desai P, et al. A systematic review on the extent and quality of pharmacoeconomic publications for China. *Value Health Region Issues* 2014;3:79–86.
- Neumann PJ, Stone PW, Chapman RH, et al. The quality of reporting in published cost-utility analyses, 1976–1997. *Ann Intern Med* 2000;132:964–72.
- Husereau D, Drummond M, Petrou S, et al. Consolidated Health Economic Evaluation Reporting Standards (CHEERS) statement. *Eur J Health Econ* 2013;14:367–72.
- Williams A. The cost-benefit approach. *Br Med Bull* 1974;30:252–6.
- Drummond MF, Jefferson TO. Guidelines for authors and peer reviewers of economic submissions to the BMJ. The BMJ Economic Evaluation Working Party. *BMJ (Clinical research ed)* 1996;313:275–83.
- Ofman JJ, Sullivan SD, Neumann PJ, et al. Examining the value and quality of health economic analyses: implications of utilizing the QHES. *J Managed Care Pharm* 2003;9:53–61.
- Chiou CF, Hay JW, Wallace JF, et al. Development and validation of a grading system for the quality of cost-effectiveness studies. *Med Care* 2003;41:32–44.
- Health Economic Evaluation Publication Guidelines (CHEERS): Good Reporting Practices [cited July 7, 2016]. Available at: <http://www.ispor.org/Health-Economic-Evaluation-Publication-CHEERS-Guidelines.asp>.
- Husereau D, Drummond M, Petrou S, et al. Consolidated Health Economic Evaluation Reporting Standards (CHEERS): explanation and elaboration: a report of the ISPOR Health Economic Evaluation Publication Guidelines Good Reporting Practices Task Force. *Value Health* 2013;16:231–50.
- Bundhun PK, Pursun M, Teeluck AR, et al. Are everolimus-eluting stents associated with better clinical outcomes compared to other drug-eluting stents in patients with type 2 diabetes mellitus? A systematic review and meta-analysis. *Medicine* 2016;95:e3276.
- Hermanowski TR, Drozdowska AK, Kowalczyk M. Institutional framework for integrated Pharmaceutical Benefits Management: results from a systematic review. *Int J Integrated Care* 2015;15:e036.
- Yu YF, Chen ZW, Zhou Z, et al. A cost-effectiveness analysis of docetaxel versus pemetrexed in second-line chemotherapy for stage IIIb or IV non-small cell lung cancer in China. *Chemotherapy* 2010;56:472–7.
- Wang S, Peng L, Li J, et al. A trial-based cost-effectiveness analysis of erlotinib alone versus platinum-based doublet chemotherapy as first-line therapy for Eastern Asian nonsquamous non-small-cell lung cancer. *PLoS One* 2013;8:e55917.
- Chen W, Jiang Z, Shao Z, et al. An economic evaluation of adjuvant trastuzumab therapy in HER2-positive early breast cancer. *Value Health* 2009;12(suppl 3):S82–4.
- Huang BT, Wang Y, Du QF, et al. Analysis of efficacy and cost-effectiveness of high-dose arabinoside versus daunorubicin chemotherapy in older adult patients with acute myeloid leukemia by cytogenetic risk profile: retrospective review from China. *Int J Hematol* 2011;93:474–81.
- Gao L, Li SC. Cost-utility analysis of liraglutide versus glimepiride as add-on to metformin in patients with type 2 diabetes in China. *Value Health* 2012;15:A663-A.
- Sun Z, Ren M, Wu Q, et al. Co-administration of Wuzhi capsules and tacrolimus in patients with idiopathic membranous nephropathy: clinical efficacy and pharmacoeconomics. *Int Urol Nephrol* 2014;46:1977–82.
- Liubao P, Xiaomin W, Chongqing T, et al. Cost-effectiveness analysis of adjuvant therapy for operable breast cancer from a Chinese perspective: doxorubicin plus cyclophosphamide versus docetaxel plus cyclophosphamide. *Pharmacoeconomics* 2009;27:873–86.
- Wu B, Shen J, Cheng H. Cost-effectiveness analysis of different rescue therapies in patients with lamivudine-resistant chronic hepatitis B in China. *BMC Health Serv Res* 2012;12:385.
- Lee KK, You JH, Wong IC, et al. Cost-effectiveness analysis of high-dose omeprazole infusion as adjuvant therapy to endoscopic treatment of bleeding peptic ulcer. *Gastrointest Endosc* 2003;57:160–4.
- Wu B, Chen H, Shen J, et al. Cost-effectiveness of adding rh-endostatin to first-line chemotherapy in patients with advanced non-small-cell lung cancer in China. *Clin Therapeut* 2011;33:1446–55.
- Palmer JL, Beaudet A, White J, et al. Cost-effectiveness of biphasic insulin aspart versus insulin glargine in patients with type 2 diabetes in China. *Adv Ther* 2010;27:814–27.
- Zeng X, Peng L, Li J, et al. Cost-effectiveness of continuation maintenance pemetrexed after cisplatin and pemetrexed chemotherapy for advanced nonsquamous non-small-cell lung cancer: estimates from

- the perspective of the Chinese health care system. *Clin Therapeut* 2013;35:54–65.
- [29] Wu B, Kun L, Liu X, et al. Cost-effectiveness of different strategies for stroke prevention in patients with atrial fibrillation in a health resource-limited setting. *Cardiovasc Drugs Ther* 2014;28:87–98.
- [30] Yang L, Li M, Tao LB, et al. Cost-effectiveness of long-acting risperidone injection versus alternative atypical antipsychotic agents in patients with schizophrenia in China. *Value Health* 2009;12(suppl 3):S66–9.
- [31] Wang M, Moran AE, Liu J, et al. Cost-effectiveness of optimal use of acute myocardial infarction treatments and impact on coronary heart disease mortality in China. *Circ Cardiovasc Qual Outcomes* 2014;7:78–85.
- [32] Chen Z, Wang C, Xu X, et al. Cost-effectiveness study comparing imatinib with interferon-alpha for patients with newly diagnosed chronic-phase (CP) chronic myeloid leukemia (CML) from the Chinese public health-care system perspective (CPHSP). *Value Health* 2009;12(suppl 3):S85–8.
- [33] Xie X, Vondeling H. Cost-utility analysis of intensive blood glucose control with metformin versus usual care in overweight type 2 diabetes mellitus patients in Beijing, P.R. China. *Value Health* 2008;11(suppl 1):S23–32.
- [34] Wu B, Ye M, Chen H, et al. Costs of trastuzumab in combination with chemotherapy for HER2-positive advanced gastric or gastroesophageal junction cancer: an economic evaluation in the Chinese context. *Clin Therapeut* 2012;34:468–79.
- [35] Chan SL, Jen J, Burke T, et al. Economic analysis of aprepitant-containing regimen to prevent chemotherapy-induced nausea and vomiting in patients receiving highly emetogenic chemotherapy in Hong Kong. *Asia-Pac J Clin Oncol* 2014;10:80–91.
- [36] Tan C, Peng L, Zeng X, et al. Economic evaluation of first-line adjuvant chemotherapies for resectable gastric cancer patients in China. *PloS One* 2013;8:e83396.
- [37] Wu B, Dong B, Xu Y, et al. Economic evaluation of first-line treatments for metastatic renal cell carcinoma: a cost-effectiveness analysis in a health resource-limited setting. *PloS One* 2012;7:e32530.
- [38] Yuan Y, Iloeje U, Li H, et al. Economic implications of entecavir treatment in suppressing viral replication in chronic hepatitis B (CHB) patients in China from a perspective of the Chinese Social Security program. *Value Health* 2008;11(suppl 1):S11–22.
- [39] Zeng X, Li J, Peng L, et al. Economic outcomes of maintenance gefitinib for locally advanced/metastatic non-small-cell lung cancer with unknown EGFR mutations: a semi-Markov model analysis. *PloS One* 2014;9:e88881.
- [40] Zhu J, Li T, Wang X, et al. Gene-guided gefitinib switch maintenance therapy for patients with advanced EGFR mutation-positive non-small cell lung cancer: an economic analysis. *BMC Cancer* 2013;13:39.
- [41] You JH, Lau W, Lee IY, et al. Helicobacter pylori eradication prior to initiation of long-term non-steroidal anti-inflammatory drug therapy in Chinese patients—a cost-effectiveness analysis. *Int J Clin Pharmacol Therapeut* 2006;44:149–53.
- [42] Gao P, Zhang H, Xu H, et al. Increased use of antidepressants in Wuhan, China: a retrospective study from 2006 to 2012. *Medicina (Kaunas, Lithuania)* 2013;49:529–34.
- [43] Zhou B, Wang J, Yan Z, et al. Liver cancer: effects, safety, and cost-effectiveness of controlled-release oxycodone for pain control after TACE. *Radiology* 2012;262:1014–21.
- [44] You JH, Lee AC, Wong SC, et al. Low-dose or standard-dose proton pump inhibitors for maintenance therapy of gastro-oesophageal reflux disease: a cost-effectiveness analysis. *Aliment Pharmacol Therapeut* 2003;17:785–92.
- [45] You JH. Novel oral anticoagulants versus warfarin therapy at various levels of anticoagulation control in atrial fibrillation: a cost-effectiveness analysis. *J Gen Intern Med* 2014;29:438–46.
- [46] Chang WC, Lee MC, Yeh LS, et al. Quality-initiated prophylactic antibiotic use in laparoscopic-assisted vaginal hysterectomy. *Austr N Z J Obstet Gynaecol* 2008;48:592–5.
- [47] Xiao YL, Nie YQ, Hou XH, et al. The efficacy, safety and cost-effectiveness of hydrotalcite versus esomeprazole in on-demand therapy of NERD: A multicenter, randomized, open-label study in China. *J Digest Dis* 2013;14:463–8.
- [48] Wei L, Hu SL, Hou J, et al. A novel estimation of the impact of treatment with entecavir on long-term mortality, morbidity, and health care costs of chronic hepatitis B in China. *Value Health Region Issues* 2013. 48–56.
- [49] Aguiar PM, Lima TM, Storpirtis S. Systematic review of the economic evaluations of novel therapeutic agents in multiple myeloma: what is the reporting quality? *J Clin Pharm Therapeut* 2016;41:189–97.
- [50] Auguste P, Tsertsivadze A, Court R, et al. A systematic review of economic models used to assess the cost-effectiveness of strategies for identifying latent tuberculosis in high-risk groups. *Tuberculosis (Edinburgh, Scotland)* 2016;99:81–91.
- [51] Snoswell C, Finnane A, Janda M, et al. Cost-effectiveness of store-and-forward teledermatology: a systematic review. *JAMA Dermatol* 2016;152:702–8.
- [52] Desai PR, Chandwani HS, Rascati KL. Assessing the quality of pharmaco-economic studies in India: a systematic review. *Pharmacoeconomics* 2012;30:749–62.
- [53] Woerschling AL, Borrego ME, Raisch DW. Assessing the quality of economic evaluations of FDA novel drug approvals: a systematic review. *Ann Pharmacother* 2016.
- [54] Gavaza P, Rascati KL, Oladapo AO, et al. The state of health economic evaluation research in Nigeria: a systematic review. *Pharmacoeconomics* 2010;28:539–53.
- [55] Jinha AE. Article 50 million: an estimate of the number of scholarly articles in existence. *Learn Publish* 2010;23:258–63.
- [56] Taddio A, Pain T, Fassos FF, et al. Quality of nonstructured and structured abstracts of original research articles in the British Medical Journal, the Canadian Medical Association Journal and the Journal of the American Medical Association. *Can Med Assoc J* 1994;150:1611–5.
- [57] Hartley J. Current findings from research on structured abstracts: an update. *J Med Library Assoc* 2014;102:146–8.