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Title: Quality assessment of systematic reviews for surgical treatment of low back pain: an overview

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1	QUALITY ASSESSMENT OF SYSTEMATIC REVIEWS
2	FOR SURGICAL TREATMENT OF LOW BACK PAIN:
3	AN OVERVIEW
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29	ABSTRACT
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21	Paskground Context: Low hash noin is smong the most frequent severe for
51	Background Context: Low back pair is among the most nequent causes for
32	medical appointments. Surgical treatment is widely controversial and new
33	surgical techniques and treatment modalities have been developed within the
34	last decade. Treatment for Low Back Pain should be evidence-based through
35	Systematic Reviews and Meta Analysis. Thus, the quality of these reviews is
36	sometimes put into question since methodological mistakes are frequently seen.

1 **Purpose**: The aim of this study is to gather all Systematic Reviews for the 2 surgical treatment of Low Back Pain and analyze their outcomes, quality and 3 conclusion. 4 **Study Design/Setting:** An overview of Systematic Reviews 5 Outcome Measures: AMSTAR score, PRISMA statement, conclusion supported 6 by descriptive statistics. 7 Methods: A literature search for Systematic Reviews containing Low Back Pain 8 surgical treatment was conducted through different medical databases. Two 9 investigators independently assessed all titles and abstracts for inclusion. Studies should have at least one surgical procedure as an intervention. Diagnoses 10 11 were categorized as lumbar disc herniation, spondylolisthesis, stenosis, facet joint syndrome, and degenerative disc disease. Quality was assessed through the 12 13 PRISMA and AMSTAR questionnaires. Study quality related to its PRISMA and/or 14 AMSTAR score percentage was rated as: very poor (<30%), poor (30-50%), fair 15 (50-70%), good (70-90%), and excellent (>90%). Articles were considered 16 conclusive if they had a conclusion for their primary outcome supported by 17 descriptive statistical evidence. This study was funded exclusively by the 18 authors' own resources. None of the authors have any potential conflict of 19 interest to declare. 20 **Results**: Overall, there were 40 systematic reviews included. According to 21 AMSTAR and PRISMA scores, 5-7.5% of the Systematic Reviews were rated as 22 excellent and most of them were rated as a fair review. AMSTAR indicated that 23 22.5% of the reviews have very poor quality, while PRISMA stated 7.5% being of 24 very poor quality. For both tools, performing a Meta Analysis made the reviews' 25 quality significantly better. The best-rated diagnosis groups according to PRISMA

1	were Spondylosis, Lumbar Disc Herniation and Degenerative Disc Disease.
2	Considering the studies' conclusions, 25 (62.5%) out of the 40 Systematic
3	Reviews had a conclusion to their primary outcome and only 11 (27.5%) were
4	supported by descriptive statistical analysis. This means that 44% of the
5	Systematic Reviews with a conclusion were evidence based. There were 15
6	(37.5%) SRs that did not reach a conclusion to their primary objectives.
7	Conclusions : In conclusion, most SRs for LBP do not reach very good or
8	excellent quality and only 27.5% of them have evidence-based conclusions.
9	Including a meta-analysis is a significant factor to improve quality and evidence
10	for SRs.
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13	Keywords: low back pain; systematic review; surgical procedures; outcome
14	assessment; lumbar spine; degenerative disc disease
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18	INTRODUCTION
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20	The impact of low back pain in healthcare is a major concern as costs are
21	increasing every year and are significantly related to surgical treatment, time to
22	return to work, and work compensation[1,2]. Surgical treatment for LBP is
23	widely controversial and new surgical techniques and treatment modalities have
24	been developed within the last decade.

1 For best scientific evidence, outcomes for new and standard interventions 2 should be analyzed through randomized clinical trials, [3-5]. However, a 3 majority of treatment modalities for LBP goes through a series of trials with 4 different outcomes amongst themselves that could be influenced by 5 methodology, population or even conflicts with authors' interests or research 6 funding.[6–9] When this happens, the best evidence for treatment outcomes is 7 derived from systematic reviews (SR) with a meta-analysis (MA) of those trials. 8 Unfortunately, there is often a great divide between those outcomes and clinical 9 practice.

10 Treatment for LBP should be evidence-based through SR and MA. 11 Systematic reviews for LBP are widely available in any medical database but they 12 might lead to different conclusions for the same intervention in the same 13 population. Thus, the quality of these reviews is sometimes put into question 14 since methodological mistakes are frequently seen[10]. Therefore, SR may not be 15 so highly evidenced to guide surgical treatment for the most common 16 degenerative lumbar spine diseases.

17 The aim of this study is to gather all SR for the surgical treatment of LBP18 and analyze their outcomes, quality and confidence.

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20 MATERIALS AND METHODS

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22 Search strategy

Institutional review board approval (number 1942-14) was obtained. A
literature search for systematic reviews that involves only LBP treatment was
conducted up to January 2014 through different medical databases: Medline

1 (Pubmed); EMBASE (Ovid); Cochrane database of systematic reviews (CDSR); 2 the Database of Abstracts of Reviews of Effectiveness (DARE); and International 3 Prospective Register of Systematic Reviews (Prospero). No restriction to 4 language or date was applied. To minimize risk of missing relevant reviews, a 5 handsearch of the reference lists of reviews captured by the initial search was 6 performed as well. The search strategy used for Medline is shown on Appendix 1. 7 Other databases followed the same search strategy with minimal adjustments. 8 Two investigators (DM and NA) independently assessed all titles and abstracts to 9 exclude duplicate articles and select potential articles to be included while inconsistencies were resolved through discussion with a third author (ML). 10 11 When more than one SR with the same or similar interventions from the same author were found, only the most current one was included and was considered 12 13 an update of previous work.

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15 Study eligibility criteria

16 After a list of studies was gathered from all database searches, systematic reviews of studies involving patients of all ages were included when the study 17 18 discussed the following diseases: lumbar disc herniation, spondylolisthesis, 19 spinal canal stenosis, facet joint syndrome and degenerative disc diseases. SRs 20 should have at least one surgical procedure as an intervention such as injections 21 of any kind, fusion, or decompression such as laminectomy or discectomy. Either 22 open or minimally invasive techniques were considered. Systematic reviews 23 comparing two non-surgical treatments were excluded as well as those involving 24 cervical or thoracic spine degenerative diseases.

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1 Data extraction

2 Data were extracted independently by four reviewers (DM, NA, MK and 3 MW) that are board certified in spine surgery with a standardized form. To 4 minimize evaluation bias, all reviewers were primarily trained for each item of 5 both questionnaires by one of the senior authors (ML) with expertise in systematic reviews. All four reviewers assessed the first 5 papers together so 6 7 there would be homogeneity on the interpretation of data. The reviewers 8 assessed the following 35 papers independently. Any disagreement that might 9 have arisen was discussed and resolved by consensus and with an opinion of a fifth reviewer (ML) with expertise in systematic reviews. The following items 10 11 were included in our form and collected for every SR: study design, searched 12 databases, last date of search strategy, presence of a protocol before conduction 13 of the study, funding sources, number of studies included, number of patients 14 assigned, number of patients assessed at the end of the study, inclusion and 15 exclusion criteria, age of participants, diagnoses enrolled, interventions, primary 16 and secondary outcomes, timing of outcome measures, and presence of positive conclusions. 17

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19 Study quality analysis

20 Quality of the studies included in the current analysis was assessed 21 through the PRISMA (Preferred Reporting Items for Systematic Reviews and 22 Meta-Analyses)[11] and AMSTAR[12] questionnaires by the same four 23 reviewers. Both forms are validated measurement tools to assess the 24 methodological quality of systematic reviews (Appendices 2 and 3). Each item of 25 the PRISMA form was graded as *yes, incomplete* or *no* and respectively scored as

1, 0.5, or 0 points for statistical analysis purposes. Similarly, the AMSTAR tool
 had each item graded as *yes* or *no* and scored as 1 or 0, respectively. The sum of
 all items scored for each questionnaire was divided by its maximum possible
 score to assess study quality as a percentage. Study quality related to its PRISMA
 and/or AMSTAR score percentage was rated as: very poor (<30%), poor (30-
 50%), fair (50-70%), good (70-90%), and excellent (>90%).

7

8 Diseases, interventions and outcomes analysis

For each SR, data were extracted for population, intervention, as well as 9 primary and secondary outcomes. Papers were grouped according to the disease 10 11 investigated (Degenerative Disk Disease (DDD), Spondylolisthesis (SL), Lumbar Stenosis (LS), Lumbar Disk Herniation (LDH), Spondylosis (S), and Facet Joint 12 13 Syndrome (FJS)) and compared. Interventions and outcomes were also compared independently. Outcomes considered were: Oswestry Disability Index 14 15 (ODI)[13], Roland Morris score[14], Short-Form 36 (SF-36)[15], visual analogic 16 scale (VAS), estimated blood loss (EBL), fusion rate, infection, operation time, 17 complications, reoperation, and return to work. Every article was reviewed by two authors to analyze the validity of the conclusion. We considered conclusive 18 19 SRs those with a conclusion for their respective primary outcome. If articles 20 were conclusive, they were further assessed for description of supportive 21 statistical evidence for this conclusion. Those that met all these criteria were 22 considered as a SR with an evidenced based conclusion.

23

24 Statistical analysis

1	Statistical analysis consisted of descriptive statistics including relative
2	and absolute frequencies. In addition, Fisher exact tests were performed to
3	compare categorical variables, and the Mann-Whitney test to compare numeric
4	variables. For all tests, a P value level less than 0.05 was considered significant.
5	All the tests have been performed using the software R 3.0.3.
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7	RESULTS
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8 9	After a full electronic search, a total of 851 references were identified
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8 9 10 11	After a full electronic search, a total of 851 references were identified (Figure 1). A title and abstract screening excluded 755 references and 49 went through a detailed eligibility process in which the full-text article was analyzed. Nine studies were excluded with reasons (Figure 1)[16–24]. Overall, there were

studies[61-66] and five studies awaiting classification[67-71] due to non-

15 English written language that were not included.

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14

17 Descriptive analysis

18 Out of the 40 included SR of surgical treatment for LBP, half of them were 19 published within the last four years[6,8,9,25,28,30,33,34,36,40,48-20 51,54,56,58,72–74] with the oldest study being from 1992[43]. The SR with the 21 greatest number of included references had 74citations[43] while there was one 22 SR with a single included study[74] (mean 17.7 studies/SR). More than half of 23 the SR (58%) had no research funding. The majority (57.5%) of the SRs did not 24 include a MA in their statistical analysis. When the diagnosis investigated was 25 considered, there were 13 SRs for degenerative disk disease, 11 for spondylosis,

six for spondylolisthesis, five for lumbar stenosis, three for lumbar disk
 herniation, and two for facet joint syndrome.

There was a very heterogeneous group of surgical interventions, including injections, direct repair of the pars, arthroplasty, decompression, nucleoplasty, endoscopic discectomy, "surgery" and fusion. There were different fusion techniques studied by the SRs, including posterolateral fusion, posterior lumbar interbody fusion, transforaminal lumbar interbody fusion, and anterior lumbar interbody fusion.

9 When outcomes were analyzed, SRs that performed a MA had most 10 significant outcomes as well as the major number of items analyzed (Table 1). 11 This comparison between outcomes from SR with a MA and without MA was 12 significant for blood loss, complications, fusion, ODI, operation time, and 13 reoperation, meaning that SR with a MA had more significant results.

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16 Descriptive study quality analysis

Percentage measures for AMSTAR, PRISMA and PRISMA domains are 17 18 shown in Table 2. To assess viability of our metrics tools for SRs quality, we have 19 conducted a comparison for correlation between AMSTAR and PRISMA through 20 a Spearman test resulting in a high level correlation coefficient (0.81; p<0.001). 21 Final AMSTAR score according to percentage indicates that most frequent rate 22 for LBP SR is in between 50 to 70%, which is a fair review. Only 7.5% of the 23 reviews are rated as excellent. 22.5% of the SRs are considered as having a very 24 poor quality. PRISMA had less reviews at the extremes of the rating measures, 25 with only 7.5% being very poor and 5% excellent. Most of the reviews were

rated as fair (37.5%). When PRISMA domains are analyzed, most flaws were
found in the abstract, methods and results sections, and these reviews tend to be
rated as very poor or poor (80% for abstract, 55% for methods, and 60% for
results). Best rates (good and excellent) are in title, introduction, discussion, and
funding domains (87.5%, 82.5%, 57.5%, and 57.5%, respectively).

6 Systematic reviews that performed a meta-analysis (n=17) were compared to those without a meta-analysis (n=23) according to their AMSTAR 7 8 and PRISMA scores. For both tools, performing a MA made the reviews' quality 9 significantly better (Figure 2). According to AMSTAR, 47% of the SR with MA was good or excellent, while 21.7% of those without MA were only good. That was 10 11 not statistically significant though (p=0.089). However, PRISMA had a 12 statistically (p=0.008) higher rate of good and excellent reviews for those where 13 a MA was performed (47.1%, versus 8.7% of those without MA). Three out of the 14 seven PRISMA domains were significantly better for SR with MA, they were 15 methods, results, and discussion (Mann-Whitney test *p*<0.001 for all three).

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17 Quality analysis according to disease studied

18 Degenerative disk disease was the most studied group with 13 SR, 19 followed by Spondylosis with 11. There were six SR for Spondylolisthesis, five 20 for Lumbar Stenosis, three for Lumbar Disc Herniation, and two for Facet Joint 21 Syndrome. The best-rated diagnosis groups according to PRISMA were 22 Spondylosis, Lumbar disk herniation and Degenerative disk disease with 36.3%, 23 33.3%, and 30.7%, respectively, for good and excellent SR (Table 3). Lumbar 24 stenosis and Facet joint disease had no SR scored over 70% according to 25 PRISMA. Furthermore, most of the papers, not considering diagnosis group, were

1 rated as very poor, poor, or fair. When the AMSTAR tool was performed, results 2 were apparently better, although not statistically significant. Half of the papers 3 within Spondylolisthesis or Facet Joint Syndrome groups were classified as good 4 or excellent, while the group with the greater number of SR rated over 70% by 5 PRISMA (Spondylosis) had the worst performance with AMSTAR (18.8% of good 6 or excellent papers). When quality of the SR was compared between diagnosis 7 groups, there was no statistical difference between either of the questionnaires' scores according to Fisher exact test. Nevertheless, SR for LDH[39,53,59] had the 8 9 smallest score for both AMSTAR and PRISMA.

10

11 Intervention and outcomes analysis

There were a wide variety of interventions. Surgery was the intervention 12 13 for all SRs since it is an inclusion criterion for this study. Specific techniques were composed of Arthroplasty (6 SRs), Fusion (10SRs), Endoscopy (1 SR), 14 15 DeKompressor®(1 SR), Injections (4 SR), Instrumentation (1 SR), Minimally 16 Invasive Interbody Fusion (2 SRs), Nucleoplasty (4 SRs), Posterolateral Fusion (4 SRs), and Osteosynthesis (1 SR). Clinical outcomes were mostly measured by the 17 18 ODI (assessed in 20 SRs), followed by the VAS pain score in 14 SRs. There were 19 five SRs analyzing SF-36 and only one used the Roland Morris score. Surgical 20 outcomes were most described as spine fusion (8 SRs), followed by EBL and 21 operation time (5 SRs each). Postoperative outcomes were measured by 22 complications (12 SRs), return to work (7 SRs), and reoperation rate (7 SRs).

Considering the studies' conclusions, 25 (62.5%) out of the 40 SRs had a conclusion to their primary outcome[6–9,27,33,34,36,38,39,42–44,48–51,53– 56,58,59,72,73] and only 11 (27.5%) were supported by descriptive statistical

1 analysis[6-9,39,48-50,56,58,73]. This means that 44% of the SRs with a 2 conclusion were evidence based. There were 15 (37.5%) SRs that did not reach a 3 conclusion to their primary objectives [25,26,28-31,35,37,40,45,52,57,60,75,76] 4 and the most frequent reason for this was the lack of available randomized 5 controlled trials. If we calculate the average PRISMA and AMSTAR scores for these 11 evidence based conclusive SRs, we see 76.6 and 75.21 respectively, 6 7 meaning that those were very good studies. If we apply the same calculus for the 8 15 SRs without a conclusion, average PRISMA and AMSTAR were respectively 9 47.28 and 41.82 meaning fair quality studies.

Among the 11 conclusive SRs, there were four studies for DDD, four for S,
one for LDH, one for LS, and one for SL. Neither of the SRs for FJS had evidence
based conclusion.

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14 **DISCUSSION**

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16 The pursuit for scientific evidence is a challenge in clinical medical. Treatments and clinical decisions should be based on the best evidence possible 17 18 but this is often not a reality due to numerous obstacles to achieve this goal. 19 Surgical treatment is often under-supported by the medical literature due to 20 heterogeneity of patients, surgical techniques, implant differences, surgeon skill, 21 and evolving new technology. The presence of statistical heterogeneity poses a 22 challenge to conduct valuable outcomes studies with reliable results that would 23 change clinical practice. Best evidence should come from systematic reviews 24 followed by meta-analysis of the data, which would ideally support all medical 25 decisions. An overload of systematic reviews has been published within the last

1 decades, with still increasing numbers on the way[77]. In our series of 40 SRs, 2 half of them were published within the last four years. Systematic reviews need 3 frequent updates as well since well-designed randomized clinical trials 4 published after a SR search was conducted may significantly change results. 5 Randomized clinical trials of the same intervention and population may disagree and SR should analyze and compare trials for definitive evidence. Although many 6 7 SRs for LBP surgical treatment are available, there is still no strong evidence 8 procedures favoring of surgical from evidence-based most an 9 approach[32,36,74,78]. Based on all literature available, spine surgeons still cannot answer most of patients' questions regarding the best treatment for 10 11 specific spine diagnoses. Searching for answers, surgeons should not blindly trust a SR since the validity of a significant number of them is questioned. Few 12 13 points should always be questioned before taking SR into practice: search conduction method; bias in the study selection; current/updated SR; quality of 14 15 the studies assessed; evidence combined and summarized appropriately; 16 publication bias; and, justified conclusions[79].

17 The main objective of this study was to assess quality of systematic 18 reviews for surgical treatment of low back pain. After 40 selected systematic 19 reviews were analyzed through the AMSTAR and PRISMA questionnaires, just a 20 few of them (7.5% and 5% respectively) were classified as excellent (scores 21 >90%). However, they were not the same according to each assessment tool. 22 PRISMA rated the works by Yajun et al^[7] and Zhou et al^[49] as excellent while 23 AMSTAR rated excellent the works by Wang et al[6], Gibson et al[39], and Jacobs 24 et al[8]. All these papers performed a meta-analysis of the data, which makes it a 25 paramount condition for a SR to be considered as excellent. Interestingly, three

1 out of the five top rated SRs had the same patient-intervention combination 2 although their results may disagree [6–8]. Not all papers elucidate their conflicts 3 of interest, which may lead to risk of bias. Those were all SRs published between 4 2010-13 comparing fusion versus total disc replacement for the degenerative 5 disc disease of the lumbar spine. Wang concludes that there is evidence that the 6 risk of clinical adjacent segment pathology following fusion is higher when 7 compared with total disc replacement. On the other hand, Yajun states that disk 8 replacement does not show significant superiority for the treatment of lumbar 9 DDD compared with fusion. Jacobs concludes that although there is no clinical relevance, total disc replacement seems to be effective in treating LBP in selected 10 11 patients and is equivalent to fusion. A fourth SR rated as fair[9] significantly favored total disc replacement for its safety and efficacy. Additionally, some 12 13 flaws in the SRs may be present but are not always detected by this study 14 metrics. For instance, although Yajuns' SR had a high score for both tools, a 15 subtle flaw was not detect, since two trials[80,81] from the same study group 16 including the same subset of patients were included and considered different papers. Even though there are already three highly qualified SRs and another 17 18 three lower scored SRs[9,57,75] on the same topic, this is still a highly debated 19 topic in every spine surgery meeting and we do not seem to be getting closer to 20 an answer. Following the conclusions above, most SRs report the need for 21 further evidence and more randomized clinical trials. In addition to conflict of 22 interest and industry role in research, caution should be taken on the 23 interpretation of industry supported or funded research while inappropriate 24 influence of funders is often regarded as an important risk of bias and it is not 25 unusual that the sponsor either owns the data or needs to approve the

1 manuscript. This is well stated in the *Cochrane Handbook for Systematic*2 *Reviews*[82]

3 Conducting an excellent SR is not an easy task, and sometimes is just not 4 feasible since after applying a well-designed search strategy and study selection, 5 authors are frequently facing insufficient data for a meta-analysis. In order to keep up with the proposed protocol, observational studies are included and 6 sometimes quality decreases. However, Furlan et al[83] stated that 7 8 observational, non-randomized studies frequently agree with randomized 9 controlled trials for low back pain. Despite all challenges, majority of the SRs for surgical treatment of LBP were considered as "good" in the present study. Those 10 11 good reviews usually have high scores for domains like title, introduction, and discussion but their biggest flaws are in the abstract, methods, and results. 12 13 According to PRISMA, this is where most studies loose their quality and authors 14 do not follow ideal patterns to conduct a SR. A significant factor for better 15 evidence in SR is applying MA to results. Studies with MA had significantly higher 16 scores than those without an MA according to both AMSTAR and PRISMA. This difference was more evident through the PRISMA tool where 47% of the SR with 17 18 a MA was rated as good or excellent against only 8.7% of those without a MA 19 (p<0.015). Many SRs for LBP achieve a final conclusion and draw benefits of a 20 technique over another. However, readers that directly access the conclusion of 21 the paper should be careful, for a significant number of SRs analyzed draw 22 conclusions without achieving evidence. Even with fair or good quality scores, 23 some SRs conclude facts that are not supported by their statistical analysis[32,35,36,38,40-42,44,54] or do not even perform a meta-analysis. For 24 25 this reason, SRs should be carefully read throughout the entire manuscript

before assumptions are taken. For example, in a SR that favors nucleoplasty for
contained disc herniation in its conclusion[54], there is no evidence or adequate
statistical analysis in its methodology and results to support conclusion. No MA
was performed as well.

5 Analyzing diagnosis groups is challenging since definition of diagnosis is not always clear. If we only take SRs that specifically analyzed LDH[39,53,59], 6 7 those had the lower mean score for both PRISMA and AMSTAR. This is not 8 significant since one the SRs in this group had the lowest score among all 9 reviews for PRISMA and significantly decreased the average in a group of three. Grouping three SRs for LDH and 13 for DDD may sound conflicting since LDH 10 11 and DDD sometimes mean the same disease. That may also be true for spondylosis, DDD, and sometimes, even lumbar stenosis. Diseases studied in SRs 12 13 for low back pain are not easy to define, and trials may gather different pain 14 generators turning the population even more heterogeneous. That happens due 15 to the clinical difficulty to define the source of pain in a patient, being all part of a 16 natural degenerative process of the spine. Two SRs analyzed facet joint 17 syndrome[38,60] and their average rate was fair reviews. Despite one of 18 them[60] was rated as good by the AMSTAR tool, its funding and conflicts of 19 interest of the authors are not very clear, which is extremely important when a 20 review of a commercial product is being made (radiofrequency). The same 21 happens in a SR of a specific percutaneous discectomy device available by the 22 spine industry[53] were funding and conflicts of interest are also questionable. 23 Spondylolisthesis may be the most non-conflicting diagnosis due to its clear 24 imaging findings. There were six SRs[28,33,34,37,50,52] for this topic with 25 average fair ratings. The most studied outcome was fusion rate for

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1 spondylolisthesis and different surgical techniques were compared. An earlier 2 study by Jacobs et al[37] in 2006 compared posterolateral fusion with other 3 interbody fusion techniques and concluded that posterolateral fusion appeared 4 to be the general gold standard for the treatment of adult isthmic 5 spondylolisthesis although there was no scientific evidence. About seven years 6 later, two SRs[33,50] rated as good comparing posterolateral fusion and 7 posterior lumbar interbody fusion for spondylolisthesis, reverted this conclusion 8 favoring interbody fusion, with significant evidence for higher fusion rates in this 9 group. The same intervention-outcome was significantly concluded by Umeta and Avanzi[48], although for a population classified as spondylosis (which 10 11 included cases with spondylolisthesis).

Although there is a trend toward minimally invasive spine surgery (MISS) 12 13 techniques, evidence is still weak. There were seven SRs concerning minimally represented by nucleoplasty[42,44,54,60], 14 invasive techniques, MISS transforaminal lumbar interbody fusion[51,76], and endoscopy techniques[59] 15 16 and none of them had significant or evidence-based conclusions. Those were 17 average, fair-quality reviews but their biggest flaw was on study selection, the 18 lack of good randomized controlled trials for these techniques made a meta-19 analysis of the data impossible. This is probably due to the late onset of these 20 techniques by spine surgeons and no good clinical trials had been performed yet.

The motivation to perform this study was the impression that most SRs for LBP surgical treatment were lacking concrete conclusions and that seemed to be a huge obstacle for SRs to have an impact on surgeons clinical routine. Our analysis proved that point when only 27.5% of the SRs for LBP surgical treatment had a conclusion for their primary objective supported by statistical

1 significance. Furthermore, 66% of the SRs relating a conclusion to their 2 objectives had no statistics to support it. If most of the SRs that could not even 3 reach a conclusion to their primary objectives put the blame on the lack of good 4 randomized clinical trials, why should we keep doing SRs if our flaw is still on 5 the primary studies? Although a good number of SRs reaches good quality scores, many of them cannot reach a conclusion due to the lack of primary 6 7 studies in the literature. An obvious increase in quality scores was observed in 8 the few SRs with a conclusion based on significant statistics while those that did 9 not determine a conclusion had the lower scores. One of the main limitations of this study was the fact that the quality tools for SRs were independently applied 10 11 by four different reviewers and inter and intra-reliability of both questionnaires 12 was not assessed. However, we have conducted a comparison for correlation 13 between AMSTAR and PRISMA through a Spearman test resulting in a high level correlation coefficient (0.81; p<0.001). Furthermore, previous authors have 14 successfully used PRISMA and AMSTAR for quality assessment of SRs of other 15 16 topics.[84-87] Another limitation is the extremely population-intervention heterogeneity of reviews which limited statistical analysis of outcomes between 17 18 SRs. The unclear definition of disease group by SRs authors may also have 19 affected analysis. An important limitation was the usage of non-validated tools to 20 determine study quality although there is none available in the literature for this 21 purpose. Future investigation of SRs for LBP is mandatory and must be constant. 22 While evidence is paramount to create treatment guidelines for LBP surgery, 23 quality analysis of the evidence provided by SRs should be assessed before 24 changing routine treatment decisions. Alongside clinical evidence, there should 25 be noted that "absence of evidence is not evidence of absence" and individual

1 interpretation should be applied to all trials and systematic reviews. Challenges 2 are always related to producing good quality randomized clinical trials, but those 3 are time consuming and bonded to high budgets. While it is easy to criticize the 4 lack of prospective, randomized studies, and those that are industry funded, 5 particularly the rigorously executed Governmental-regulated trials, funding is 6 almost an indispensable condition for a large well-performed randomized 7 clinical trial. In our opinion, we should not ban funding, but prioritize well-8 design studies proposals following strict methodological validated standards 9 even before the trial starts. Protocols should be registered and published, before started, in an international peer-reviewed periodic. A clear well-performed 10 11 methodology would minimize funding bias.

In conclusion, most systematic reviews for low back pain do not reach very good or excellent quality and only 27.5% of them have evidence-based conclusions. Therefore, authors suggest that researchers should concentrate efforts in performing randomized clinical trials in surgical treatment for low back pain before attempting secondary studies. Including a meta-analysis is a significant factor to improve quality and evidence for systematic reviews.

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Figure Legends:

- Fig. 1 Flow chart for study screening and eligibility.
- Fig. 2 Boxplots for AMSTAR and PRISMA total percentage according to systematic reviews (SR) with or without meta-analysis (MA). Mann-Whitney test.

Outcomes	Results	SR with MA (n=17)	SR without MA (n=23)	p-value
		n (%)	n (%)	
	NA	12 (70.6)	23 (100)	
Estimated blood	NS	3 (17.6)	0 (0)	0 009
loss	SR	2 (11.8)	0 (0)	0.005
	NA	8 (47.1)	21 (91.3)	
Complications	NS	6 (35.3)	2 (8.7)	0.004
complications	SR	3 (17.6)	0(0)	0.001
	NA	9 (52.9)	23 (100)	
Fusion	NS	3 (17.6)	0 (0)	<0.001
	SR	5 (29.4)	0 (0)	
	NA	16 (94.1)	23 (100)	
Infection	NS	1 (5.9)	0 (0)	0.425
	NA	5 (29.4)	15 (65.2)	
ODI	NS	7 (41.2)	7 (30.4)	0.035
001	SR	5 (29.4)	1 (4.3)	
	NA	12 (70.6)	23 (100)	
Operation time	NS	2 (11.8)	0 (0)	0.009
	SR	3 (17.6)	0 (0)	
	NA	9 (52.9)	17 (73.9)	
Pain VAS	NS	6 (35.3)	4 (17.4)	0.385
	SR	2 (11.8)	2 (8.7)	
	NA	10 (58.8)	23 (100)	
Reoperation	NS	5 (29.4)	0 (0)	0.001
·	SR	2 (11.8)	0 (0)	
	NA	14 (82.4)	19 (82.6)	
Return to work	NS	2 (11.8)	3 (13)	1
	SR	1 (5.9)	1 (4.3)	

Roland-Morris	NA	17 (100) 0 (0)	22 (95.7) 1 (4 3)	1
65.26	NA	14 (82.4)	21 (91.3)	0.024
SF-36	NS	3 (17.6)	2 (8.7)	0.634

Table 1 – Analysis of outcomes considering systematic reviews with or without a meta-analysis. SR = systematic review; MA =meta-analysis; NA = not analyzed in the systematic review; NS = non-significant; SR = significant result; ODI = Oswestry disability index; VAS = visual analogic scale; SF-36 = Short-Form 36. *p* values are from Fisher exact test.

Quality tool		С	ategories	×	
	Very Poor	Poor	Fair	Good	Excelent
	%	%	%	%	%
Amstar total	22.5	15	30	25	7.5
Prisma total	7.5	30	37.5	20	5
Prisma - Title	12.5	0	0	0	87.5
Prisma - Abstract	17.5	62,5	0	0	20
Prisma - Introduction	0	17.5	0	37.5	45
Prisma - Methods	22.5	32.5	20	20	5
Prisma - Results	35	25	12.5	15	12.5
Prisma - Discussion	5	17.5	20	7.5	50
Prisma - Funding	37.5	5	0	0	57.5

Table 2 - AMSTAR and PRISMA quality distribution by percentage of final score. PRISMA is alsosubdivided by domains percentages. Very poor (<30%); Poor (30-50%); Fair (50-70%); Good (70-</td>90%); Excellent (>90%)

C			
	п	PRISMA >70	AMSTAR >70
DDD	13	4 (30.7%)	5 (38.4%)
SL	6	1 (16.6%)	3 (50%)
LS	5	0 (0)	1 (20%)
LDH	3	1 (33.3%)	1 (33.3%)
S	11	4 (36.3%)	2 (18.8%)
FJS	2	0 (0)	1 (50%)

Table 3 – Absolute and relative frequencies of very good and excellent systematic reviews according to disease group in PRISMA and AMSTAR tools. DDD = Degenerative disc disease, SL = Spondylolisthesis, LS = Lumbar stenosis, LDH = Lumbar disc herniation, S = Spondylosis, FJS = Facet joint syndrome. P=0.713 (PRISMA) and p=0.747 (AMSTAR). Fisher exact test.

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