

Clinical practice guidelines for the noninvasive management of low back pain: A systematic review by the Ontario Protocol for Traffic Injury Management (OPTIMa) Collaboration

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Conflicts of interest

Dr. Côté reports a grant from the Ontario Ministry of Finance, and funding from the Canada Research Chairs Program - Canadian Institutes of Health Research (CIHR) during the conduct of the study; expert testimony for Canadian Chiropractic Protective Association; payment for lectures and other expenses from the European Spine Society outside the submitted work. Dr. Gross

Abstract

We conducted a systematic review of guidelines on the management of low back pain (LBP) to assess their methodological quality and guide care. We synthesized guidelines on the management of LBP published from 2005 to 2014 following best evidence synthesis principles. We searched MEDLINE, EMBASE, CINAHL, PsycINFO, Cochrane, DARE, National Health Services Economic Evaluation Database, Health Technology Assessment Database, Index to Chiropractic Literature and grev literature. Independent reviewers critically appraised eligible guidelines using AGREE II criteria. We screened 2504 citations; 13 guidelines were eligible for critical appraisal, and 10 had a low risk of bias. According to highquality guidelines: (1) all patients with acute or chronic LBP should receive education, reassurance and instruction on self-management options; (2) patients with acute LBP should be encouraged to return to activity and may benefit from paracetamol, nonsteroidal antiinflammatory drugs (NSAIDs), or spinal manipulation; (3) the management of chronic LBP may include exercise, paracetamol or NSAIDs, manual therapy, acupuncture, and multimodal rehabilitation (combined physical and psychological treatment); and (4) patients with lumbar disc herniation with radiculopathy may benefit from spinal manipulation. Ten guidelines were of high methodological quality, but updating and some methodological improvements are needed. Overall, most guidelines target nonspecific LBP and recommend education, staying active/exercise, manual therapy, and paracetamol or NSAIDs as first-line reports honorarium and support for travel to meetings from the University of Ontario Institute of Technology during the conduct of the study; research operating grants from the workers' compensation boards of Alberta, Manitoba and WorkSafe BC outside the submitted work. Dr. Carroll reports a grant from CIHR outside the submitted work. Dr. Nordin reports support for travel to meetings for the Minor Injury Guideline during the conduct of this study. Dr. Mior reports honorarium from University of Ontario Institute of Technology during the conduct of the study; fees for consultancy from Ontario Chiropractic Association, grants from Ontario Chiropractic Association, payment for lectures from Manitoba Workers Compensation Board, payment for manuscript preparation and other expenses from Canadian Memorial Chiropractic College, outside the submitted work. For the remaining authors, no conflicts were declared.

Database: MEDLINE, EMBASE, CINAHL, Psy-CINFO, Cochrane Database of Systematic Reviews, Database of Abstracts of Reviews of Effects, Cochrane Central Register of Controlled Trials, National Health Services Economic Evaluation Database, Health Technology Assessment Database, the Index to Chiropractic Literature and the grey literature.

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1. Introduction

More than 80% of people experience at least one episode of back pain during their lifetime (Cassidy et al., 1998; Walker, 2000). Back pain is a common source of disability, whether the pain is attributed to work, traffic collisions, activities of daily living, or insidious onset (Cassidy et al., 1998, 2005; Hincapie et al., 2010). Back pain is costly, accounting for a considerable proportion of work absenteeism and lost productivity (Carey et al., 1995, 1996). Moreover, it is the most common reason for visiting a healthcare provider for musculoskeletal complaints (Cypress, 1983; Côté et al., 2001). Although multiple clinical interventions are available to treat back pain, current

treatments. The recommendation to use paracetamol for acute LBP is challenged by recent evidence and needs to be revisited.

Significance: Most high-quality guidelines recommend education, staying active/exercise, manual therapy and paracetamol/NSAIDs as first-line treatments for LBP. Recommendation of paracetamol for acute LBP is challenged by recent evidence and needs updating.

evidence suggests that their effects appear small and short term (Haldeman and Dagenais, 2008).

Clinical practice guidelines are systematically developed statements that include recommendations intended to optimize patient care and improve patients' health outcomes (Shekelle et al., 1999, 2012; Institute of Medicine Committee on Standards for Developing Trustworthy Clinical Practice Guidelines, 2011). Guidelines aim to reduce the gap between research and clinical practice and assist policy makers with decisions that impact the population (Whitworth, 2006; Alonso-Coello et al., 2010). However, concerns have been raised about the quality of many clinical practice guidelines (Ransohoff et al., 2013). Systematic reviews report that some guidelines have methodological limitations (Shaneyfelt et al., 1999; Graham et al., 2001; Hasenfeld and Shekelle, 2003; Alonso-Coello et al., 2010; Berrigan et al., 2011; Knai et al., 2012). Common flaws include poor literature review methodology, limited involvement of stakeholders and unclear editorial independence (Alonso-Coello et al., 2010). Therefore, valid concerns exist about the potentially negative impact of biased guidelines on the care and health outcomes of patients (Delgado-Noguera et al., 2009; Shaneyfelt and Centor, 2009; Tricoci et al., 2009; Alonso-Coello et al., 2010).

Guidelines of poor methodological quality may lead clinicians to consider interventions that are ineffective, costly, or harmful. Low-quality guidelines may lead decision makers to invest in the implementation of ill-informed recommendations. Moreover, low-quality guidelines may reduce their adoption by clinicians and policy makers. Known barriers to the adoption of guidelines include lack of clarity of recommendation development, ambiguous recommendations, and inconsistent recommendations across guidelines (Cote et al., 2009). Finally, when combined with other barriers, such as lack of time, limited understanding of how guidelines are developed, and inadequate dissemination, it is easy to understand why the uptake of some clinical guidelines by clinicians has been disappointing (Cote et al., 2009; Bishop et al., 2015; Slade et al., 2015).

Many clinical practice guidelines on the management of low back pain are available in the peerreviewed literature. A systematic review of these guidelines found that the quality of their methodology was adequate but varied across guidelines (Dagenais et al., 2010). However, the literature search for this systematic review ended in 2009 (Dagenais et al., 2010), and many guidelines have been published or updated since (Cutforth et al., 2011; Livingston et al., 2011; Philippine Academy of Rehabilitation Medicine. 2011: Brosseau et al., 2012: Delitto et al., 2012; Kung et al., 2012; North American Spine Society, 2012; Scottish Intercollegiate Guidelines Network, 2013; Kreiner et al., 2014). An up-to-date systematic review of these guidelines is needed to assess their methodological quality and help guide appropriate management of low back pain.

The purpose of this systematic review was to review clinical practice guidelines, programmes of care, and treatment protocols to identify effective conservative (noninvasive) interventions for the management of acute and chronic low back pain.

2. Methods

2.1 Review registration

The protocol for our systematic review was registered on PROSPERO (CRD42015017762) and can be accessed at www.crd.york.ac.uk/PROSPERO/display_ record.asp?ID=CRD42015017762.

2.2 Literature search

We developed the search strategy in consultation with a health sciences librarian. A second librarian reviewed the search strategy using the Peer Review of Electronic Search Strategies Checklist (Sampson et al., 2009). The search strategy combined terms relevant to low back pain and guidelines and included free-text words and subject headings specific to each database (Supporting Information Appendix S1). The following databases were searched from January 1, 2005, to April 30, 2014: MEDLINE, EMBASE, CINAHL, PsycINFO, Cochrane Database of Systematic Reviews, Database of Abstracts of Reviews of Effects, Cochrane Central Register of Controlled Trials, National Health Services Economic Evaluation Database, Health Technology Assessment Database and the Index to Chiropractic Literature. Guidelines published prior to 2005 were considered outdated (Kung et al., 2012) and were captured in a previous systematic review of guidelines (Dagenais et al., 2010). We hand searched reference lists of relevant guidelines for supplemental documents relevant to the methodology of that guideline.

We searched the grey literature using the following: National Guideline Clearinghouse (Agency for Healthcare Research and Quality), Canadian Medical Association Infobase, Guidelines International Network, PEDro, Trip Database, American College of Physicians Clinical Recommendations, Australian Government, National Health and Medical Research Council, Health Services/Technology Assessment Texts, Institute for Clinical Systems Improvement, National Institute for Health and Clinical Excellence (NICE) Guidance, NICE Pathways, New Zealand Guidelines Group, Scottish Intercollegiate Guidelines Network (SIGN), and World Health Organization guidelines approved by the Guidelines Review Committee.

2.3 Study selection

We used the following inclusion criteria: (1) English language; (2) targeting adults and/or children with low back pain with or without radiculopathy; (3)

guidelines, programmes of care, or treatment protocols; (4) including recommendations for therapeutic noninvasive management.

We excluded guidelines that: (1) did not include treatment recommendations; (2) were a summary or copy of previous guidelines; (3) were developed solely on the basis of consensus opinion; (4) did not conduct a systematic literature search or critical appraisal of studies used to derive recommendations; and (5) only targeted invasive (e.g. injection, surgery) interventions.

2.4 Title and abstract screening

We used a two-stage (title/abstracts and full-text) screening process with random pairs of independent reviewers. Disagreements between pairs of reviewers were resolved by discussion. A third reviewer was used to resolve disagreements if consensus could not be reached. We contacted authors if additional information was necessary to determine eligibility.

2.5 Critical appraisal of eligible guidelines

Randomly allocated pairs of independent reviewers appraised relevant guidelines using the Appraisal of Guidelines for Research and Evaluation II (AGREE II) instrument (Table 1; Brouwers et al., 2010). The AGREE II instrument is widely used to assess the development and reporting of guidelines. It consists of 23 items in six quality-related domains: scope and purpose, stakeholder involvement, rigour of development, clarity of presentation, applicability, and editorial independence of guidelines (Table 1). All reviewers were trained in critical appraisal of guidelines using the AGREE II instrument. Discussions were held between paired reviewers to reach consensus on: (1) individual AGREE II items; (2) overall guideline guality; (3) whether the guideline was high quality; and (4) whether modifications to the guideline would be needed for use in specific jurisdictions (e.g. updating literature, modifying the format of the guideline). contacted additional We authors if information was needed to complete the critical appraisal.

Guidelines with poorly conducted systematic literature searches (question 7 of AGREE II) or with inadequate methods to critically appraise the evidence (question 9 of AGREE II) were deemed to have fatal flaws and were excluded from our synthesis. These criteria are described as fundamental steps to the development of evidence-based guidelines Table 1 The AGREE II instrument (Brouwers et al., 2010).

AGREE II domains and items

Domain 1. Scope and purpose

- 1. The overall objective(s) of the guideline is (are) specifically described.
- 2. The health question(s) covered by the guideline is (are) specifically described.
- The population (patients, public, etc.) to whom the guideline is meant to apply is specifically described.

Domain 2. Stakeholder involvement

- 4. The guideline development group includes individuals from all the relevant professional groups.
- 5. The views and preferences of the target population (patients, public, etc.) have been sought.
- 6. The target users of the guideline are clearly defined.
- Domain 3. Rigour of development
- 7. Systematic methods were used to search for evidence.
- 8. The criteria for selecting the evidence are clearly described.
- 9. The strengths and limitations of the body of evidence are clearly described.
- 10. The methods for formulating the recommendations are clearly described.
- 11. The health benefits, side-effects, and risks have been considered in formulating the recommendations.
- 12. There is an explicit link between the recommendations and the supporting evidence.
- 13. The guideline has been externally reviewed by experts prior to its publication.
- 14. A procedure for updating the guideline is provided.
- Domain 4. Clarity of presentation
- 15. The recommendations are specific and unambiguous.
- 16. The different options for management of the condition or health issue are clearly presented.
- 17. Key recommendations are easily identifiable.

Domain 5. Applicability

- 18. The guideline provides advice and/or tools on how the recommendations can be put into practice.
- 19. The guideline describes facilitators and barriers to its application.
- 20. The potential resource implications of applying the
- recommendations have been considered.
- 21. The guideline presents monitoring and/or auditing criteria.
- Domain 6. Editorial independence
- 22. The views of the funding body have not influenced the content of the guideline.
- Competing interests of guideline development group members have been recorded and addressed.

AGREE II, Appraisal of Guidelines for Research and Evaluation, Version II.

(Ransohoff, 2013). Although not considered a fatal flaw, we considered lack of editorial independence from the funding body (question 22 of AGREE II) an important limitation to the quality of the guideline. The absence of editorial independence would contribute to lower overall guideline quality, since this may suggest poor reporting and lack of transparency in guideline development (Alonso-Coello et al., 2010).

2.6 Data extraction

One reviewer extracted data from high-quality guidelines and built evidence tables. A second reviewer checked the data that were extracted from each guideline by comparing the extracted data with the data reported in the guidelines. We did not extract data on the use of interventional (invasive, surgical) therapies.

2.7 Data synthesis

We synthesized recommendations from high-quality guidelines using evidence tables. Recommendations from high-quality guidelines were synthesized by interventions and summarized according to whether an intervention is (1) recommended; (2) not recommended or (3) lacked evidence to support or refute its use. We considered an intervention to be 'recommended' if the high-quality guideline used the following terminology: 'strongly recommended', 'recommended without any conditions required', 'should be used', or 'recommended for consideration' [includes 'offer' or 'consider' (National Institute of Health and Care Excellence, 2014)]. We stratified recommendations by duration of low back pain (i.e., acute or chronic) and by the number of guidelines recommending the intervention ('recommended by all guidelines' or 'recommended by most guidelines', i.e., more than 50% of guidelines).

3. Results

We screened 2504 titles and abstracts for eligibility (Fig. 1). Of those, 75 potentially relevant articles were assessed in full-text screening and 61 were ineligible. Primary reasons for ineligibility during full-text screening were (1) no systematic search or critical appraisal methods (8/61); (2) ineligible study design (48/61): (3) ineligible interventions (4/61): and (4) ineligible population (1/61). We critically appraised 13 eligible guidelines (reported in 14 articles/publications) and needed to contact authors of five guidelines (3/5 responded) to obtain additional information to assess guideline quality (Airaksinen et al., 2006; Nielens et al., 2006; Livingston et al., 2011). We identified 10 high-quality guidelines (Airaksinen et al., 2006; Nielens et al., 2006; van Tulder et al., 2006; Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011; Delitto et al.,



Figure 1 Flow diagram of the selection of guidelines on the management of low back pain.

2012; North American Spine Society, 2012; Scottish Intercollegiate Guidelines Network, 2013; Kreiner et al., 2014). Inter-rater agreement for article screening was k = 0.66 (95% confidence intervals 0.51; 0.81). Percentage agreement for guideline admissibility during independent critical appraisal was 77% (10/13). We reached consensus through discussion for the three guidelines where there was disagreement between reviewers' independent appraisal review (Livingston et al., 2011; Philippine Academy of Rehabilitation Medicine, 2011; Brosseau et al., 2012).

3.1 Methodological quality

The methodological quality of the 13 relevant guidelines varied (Tables 2 and 3). Most guidelines did not adequately address guideline applicability, particularly facilitators and barriers, resource implication, and/or monitoring or auditing criteria upon implementation (8/13 guidelines; Airaksinen et al., 2006; Nielens et al., 2006; Chou et al., 2007, 2009; Livingston et al., 2011; Brosseau et al., 2012; Delitto et al., 2012; Kreiner et al., 2014). Similarly, most guidelines did not clearly indicate whether they sought the views or preferences of the target population (9/13 guidelines; Airaksinen et al., 2006; van Tulder et al., 2006; Chou et al., 2007, 2009; Cutforth et al., 2011; Livingston et al., 2011; Brosseau et al., 2012; Delitto et al., 2012; Kreiner et al., 2014).

The 10 guidelines with high methodological quality met the following criteria: (1) systematic methods to search for evidence (10/10); (2) clearly described strengths and limitations of the evidence (10/10); (3) considered health benefits, side-effects and risks (10/ 10); (4) provided an explicit link between recommendations and supporting evidence (10/10); (5) clearly described methods for formulating recommendations (9/10); and (6) clearly described criteria for selecting evidence (7/10; Table 2). However, the high-quality guidelines had limitations, including (1) no description of an external review process (5/10) (Airaksinen et al., 2006; Nielens et al., 2006; van Tulder et al., 2006; Chou et al., 2007; Livingston et al., 2011); (2) no description of the procedure to update the guideline (3/10) (Airaksinen et al., 2006; Nielens et al., 2006; van Tulder et al., 2006); or (3) no declaration of competing interests by the guideline development group (2/10) (Airaksinen et al., 2006; Livingston et al., 2011). Six guidelines were published more than 5 years ago and need to be updated (Airaksinen et al., 2006; Nielens et al., 2006; van Tulder et al., 2006; Chou et al., 2007,

2009; National Institute of Health and Care Excellence, 2009).

The three low-quality guidelines had major limitations: (1) no clear selection criteria of the literature (2/3: Philippine Academy of Rehabilitation Medicine. 2011; Delitto et al., 2012); (2) no clear description of strengths and limitations of the literature (2/3; Brosseau et al., 2012; Delitto et al., 2012); (3) no clear description of the methods used to formulate recommendations (3/3; Philippine Academy of Rehabilitation Medicine, 2011; Brosseau et al., 2012; Delitto et al., 2012); (4) no description of side-effects and risks (2/3; Philippine Academy of Rehabilitation Medicine, 2011; Brosseau et al., 2012); (5) no description of editorial independence from funders (1/3; Delitto et al., 2012); and (6) no declaration of whether there were any competing interests by guideline development group (3/3; Philippine Academy of Rehabilitation Medicine, 2011; Brosseau et al., 2012; Delitto et al., 2012).

3.2 High-quality guidelines

Nine of the 10 high-quality guidelines addressed nonspecific low back pain (Table S1 and Table 4). Of these, one guideline targeted acute low back pain (van Tulder et al., 2006), five targeted chronic low back pain (Airaksinen et al., 2006; Nielens et al., 2006; Chou et al., 2009; National Institute of Health and Care Excellence, 2009; Scottish Intercollegiate Guidelines Network, 2013), and three addressed both acute and chronic (Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011). For chronic low back pain, one guideline commented on multimodal rehabilitation (combined physical and psychological interventions) only (i.e., no recommendations for any other noninvasive interventions; Chou et al., 2009). The remaining guideline targeted lumbar disc herniation with radiculopathy (Table S1 and Table 4; Kreiner et al., 2014).

3.3 Acute nonspecific low back pain (four highquality guidelines)

Interventions recommended by all guidelines:

- (1) Advice, reassurance, or education with evidencebased information on expected course of recovery and effective self-care options for pain management (van Tulder et al., 2006; Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011).
- (2) Early return to activities, staying active, or avoiding prescribed bed rest (van Tulder et al., 2006; Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011).

(a) Items 1–12					2								
					Views of		Systematic		Strengths	Methods for	Benefits,	Link b	letween
GDG, Year	Overall objectives	Health questions	Population	GDG	target population	Target users	search methods	Selection criteria	and limitations of evidence	formulating recommenda	side-effec tions risks	tts, recom and e	nmendations vidence
Oregon Health Authority, 2011 (Livineston et al., 2011)	7	4	2	Ŷ	_	7	4	P	~	9	2	ъ	
NICE, 2009 (NICE, 2009)	7	9	7	9	9	5	7	7	9	IJ	7	5	
North American Spine Society,	7	5	6	4	1	7	7	1	ß	6	4	7	
2012 (Kreiner et al., 2014)													
COST B13 Working Group, 2006a	7	7	7	7	1	2	5	2	D	D	9	7	
(acute) (van Tulder et al., 2006)													
COST B13 Working Group, 2006b	7	6	7	5	1	5	5	9	4	5	9	7	
(chronic) (Airaksinen et al., 2006)													
Alberta, 2012 (Cutforth et al., 2011)	7	7	7	7	1	7	7	7	7	6	9	2	
American Pain Society, 2009	6	7	7	4	1	4	S	5	6	S	7	7	
(Chou et al., 2009)													
American College of Physicians and	6	6	6	с	2	6	4	1	ы	7	9	7	
the American Pain Society, 2007													
(Chou et al., 2007)													
Belgian Health Care Knowledge	5	6	7	2	1	2	7	7	7	2	7	7	
Centre, 2006 (Nielens, 2006)													
SIGN, 2013 (Scottish Intercollegiate	7	7	7	ý	9	7	7	5	10	2	7	9	
Guidelines Network, 2013)													
(b) Items 13–23													
												Views	
												of the	Competing
	External	Procedure	Specific an	dations	Options	Key		Facilitators	Advice	Resource	Monitoring and	funding	interests
uuu, tear	review	IOF Upuduirig	Leconnine	SUNDE	preserieu	Lecul	IITIETIdauuris	driu barrier		ITTIPIICAUUTI	audiung criteria	pody	01 60 6

GDG, Year	External review	Procedure for updating	Specific and clear recommendations	Options presented	Key recommendations	Facilitators and barriers	Advice and tools	Resource implication	Monitoring and auditing criteria	of the funding body	Competing interests of GDG
Oregon Health Authority, 2011 (Livingston et al., 2011)	2	7	7	2	7	-	m	2	1	_	_
NICE, 2009 (NICE, 2009)	5	7	5	7	7	5	9	7	ŝ	7	7
North American Spine Society,	5	7	6	6	6	1	1	c.	1	5	7
2012 (Kreiner et al., 2014)											
COST B13 Working Group, 2006a	-	с	7	7	7	6	6	1	1	1 ^b	6
(acute) (van Tulder et al., 2006)											
COST B13 Working Group, 2006b	2	1	7	7	7	1	1	4	2	7	7
(chronic) (Airaksinen et al., 2006)											
Alberta, 2012 (Cutforth et al., 2011)	5	6	7	7	7	7	7	5	1	7	7

Table 2 (Continued)											
(b) Items 13–23											
GDG, Year	External review	Procedure for updating	Specific and clear recommendations	Options presented	Key recommendations	Facilitators and barriers	Advice and tools	Resource implication	Monitoring and auditing criteria	Views of the funding body	Competing interests of GDG
American Pain Society, 2009 (Chou et al. 2000	4	4	7	7	7	ε	-	-	-	7	Q
American College of Physicians and the American Pain Society, 2007	2	4	7	6	7		с		F	2	4
(Chou et al., 2007) Belgian Health Care Knowledge Centre 2006 (Nielens 2006)	~	-	ы	2	4	-	-		-	7	7
SIGN, 2013 (Scottish Intercollegiate Guidelines Network, 2013)	2	9	7	7	7	4	Ŷ	IJ	Q	7	4
AGREE II, Appraisal of Guidelines Guidelines Network. ^a Each AGREE II item is rated on a	for Resea	rrch and Evaluati tale, where 1 = 1	ion, Version II; GDG, strongly disagree or i	Guideline Dev insufficient int	velopment Group; Nil formation provided a	CE, National Ins ind 7 = strongly	stitute of Hea / agree.	alth and Care	Excellence; SIGN, S	Scottish In	tercollegiate

- (3) Paracetamol (acetaminophen) or nonsteroidal anti-inflammatory drugs (NSAIDs) if indicated (van Tulder et al., 2006; Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011), with advice and consideration of risks and warning symptoms and signs associated with these medications. Only one guideline specified the recommended type and dosage of NSAID use [i.e. Ibuprofen, up to 800 mg three times per day (maximum of 800 mg four times per day) or diclofenac, up to 50 mg three times per day] (Cutforth et al., 2011).
- (4) Muscle relaxants (short course) alone or in addition to NSAIDs if an initial trial of paracetamol or NSAIDs failed to reduce pain on their own (van Tulder et al., 2006; Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011), with advice and consideration of sedation risks associated with muscle relaxants (Chou et al., 2007; Livingston et al., 2011). Only one guideline specified the recommended type and dosage of muscle relaxant use (i.e. Cyclobenzaprine, 10–30 mg/day, with greatest benefit within 1 week, although up to 2 weeks may be justified) (Cutforth et al., 2011).
- (5) Spinal manipulation for those not improving with self-care options (Chou et al., 2007; Liv-ingston et al., 2011) or failing to return to normal activities (van Tulder et al., 2006; Cutforth et al., 2011).

Interventions recommended by most guidelines:

 Short-term use of opioids on rare occasions, to control refractory, severe pain (3/4 guidelines) (Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011). However, long-term use of opioids may be associated with significant risks related to the potential for tolerance, addiction or abuse (Livingston et al., 2011). One guideline did not address opioids for acute low back pain (van Tulder et al., 2006).

3.4 Chronic nonspecific low back pain (eight high-quality guidelines)

Interventions recommended by all guidelines:

(1) Education including advice and information promoting self-management (Cutforth et al., 2011); evidence-based information on expected course and effective self-care options (Chou et al., 2007; Livingston et al., 2011); brief educational interventions for short-term improvement (Airaksinen et al., 2006); and advice to stay active or make an early return to activities as tolerated

²Rating is based on current information available.

(a) Items 1–12													
GDG, Year	Overall objectives	Health questions	Population	GDG	Views of target population	Target users	Systematic search methods	Selection	strengths and imitations of evidence	Methods for formulating recommendatio	Benefits, side-effects, ns risks	Link betwee recomr and ev	en mendations idence
Ottawa Panel, 2012 (Brosseau et al 2012)	9	9	7	9	_	7	5	9	4	б	2	ъ	
American Physical Therapy Association, 2012 (Delitto	7	D	Ŀ	9	, -	7	m	~		-	Ŀſ	D	
et al., 2012) Philippine Academy of Rehabilitation Medicine, 2011 (Philippine Academy of Rehabilitation Medicine, 2011)	М	4	4	ы	-	Ŷ	ъ	N	N	4	0	4	
(b) Items 13–23													
GDG. Year	External review	Procedure for updating	Specific and clear recommen	dations	Options presented	Key	mendations	Facilitators and barriers	Advice and tools	Resource Mi implication au	onitoring and diting criteria	/iews of the unding	Competing interests of GDG
Ottawa Panel, 2012 (Brosseau	5		5		. 7	-		-	-				_
et al., 2012/ American Physical Therapy Association, 2012 (Delitto et al. 2012)	Ъ	Q	6		7	~			Ν	-		_	~-
Philippine Academy of Rehabilitation Medicine, 2011 (Philippine Academy of Rehabilitation Medicine, 2011)	~	4	Q		Q	Ъ		ы	4	2			-
AGREE II, Appraisal of Guideline: ^a Each AGREE II item is rated on	for Researc a 7-point sca	th and Evaluaties, where 1 =	cion, Version I strongly dise	II; GDG, agree ol	Guideline Dev insufficient i	/elopmer nformatic	nt Group. on provided a	ind 7 = stron	gly agree.				

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Table 3 AGREE II ratings^a of low-quality guidelines: (a) Items 1-12 and (b) Items 13-23.

			Early										
		Advice,	return to						Muscle				
		education,	activities				Passive		relaxant		Opioid		
Guideline	Duration of Low Back Pain	self-management or reassurance	or staying active	Exercise	Manual therapy	Acupuncture	physical modality	Acetaminophen or NSAID	(short course)	Gabapentin	(short course)	Antidepressant	Multimodal rehabilitation
COST R13 Working	Acrite	۵	<u>م</u>		<u>م</u>			۵	α.				
Group, 2006 (Airaksinen	Chronic	: œ		2	: @		RA	: 22	R ^c (cautioned	RA	Rc	R ^c	2
et al., 2006; van									against				
Tulder et al., 2006)									long-term				
									use due to				
									side-effects)				
Belgian Health Care	Chronic	Ж	Ж	Я	Я	Я	RA	Я	R ^c (cautioned	RA	Rc		н
Knowledge Centre,	(>12 weeks)								against				
2006 (Nielens, 2006)									long-term				
									use due to				
									side-effects)				
American College of	Acute	£	ш		Ы			ĸ	Я		R ^c		
Physicians and the	Chronic	с	Я	Ж	Я	Я	RA	Ж	RA		Rc	R ^c	Я
American Pain Society,													
2007 (Chou et al., 2007)													
American Pain Society,	Chronic												Ч
2009 ^b													
(Chou et al., 2009)													
NICE, 2009 (National	Chronic	۲	Ж	Ж	Ж	Ж	RA	ш			Rc	R ^c	ы
Institute of Health and	(6 weeks to												
Care Excellence, 2009)	12 months)												
Oregon Health Authority,	Acute	Ч	Я		Ж			Я	Я		Rc		
2011 (Livingston et al.,	Chronic	ч	ш	Ж	Ж	Я		Я	RA		Rc	R ^c	Ы
2011)													
Institute of Health	Acute and	ч	ш		Ж			Я	Я		Rc		
Excellence, 2012	subacute												
(Cutforth et al., 2011)	(≤12 weeks)												
	Chronic	Ъ	Ч	Ч	Ы	Ы	RA	Я	RA	RA	R ^c	R ^c	Ы
	(>12 weeks)												
SIGN, 2013 (Scottish	Chronic	Ч	Ч	ж	ы	Я	R (for laser	Я	RA	RA	Rc	RA	Я
Intercollegiate Guidelines	(>12 weeks)						only based on						
Network, 2013)							inconsistent						
							evidence)						
NICE. National Institute	of Health and	Clinical Excellence	e; NSAID, no	n-steroida	I anti-inflé	ammatory dru	igs; SIGN, Scotti	ish Intercollegiate	e Guidelines Ne	etwork; R, re	ecommen	ded (includes i	nterventions

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that are strongly recommended or recommended for consideration); RA, recommended against (includes interventions that should not be offered).

aRecommendations that were not specific to the duration of low back pain were presumed to apply to both acute and chronic.

 $^{\mathrm{b}\mathrm{T}}$ his guideline focused on the effectiveness of multimodal rehabilitation for chronic low back pain.

 $^{\circ}$ These guidelines recommended considering significant risks and possible side-effects.

(Airaksinen et al., 2006; Nielens et al., 2006; Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013).

- (2) Exercises (Nielens et al., 2006; Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013) including supervised exercises (Airaksinen et al., 2006; National Institute of Health and Care Excellence, 2009) or yoga (Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011). Three guidelines found insufficient evidence to make recommendations for or against any specific type of exercise (Airaksinen et al., 2006; Nielens et al., 2006; Scottish Intercollegiate Guidelines Network, 2013), but to instead consider patient preferences (Airaksinen et al., 2006). Recommended frequency/duration was a maximum of eight sessions over up to 12 weeks (National Institute of Health and Care Excellence, 2009).
- (3) Manual therapy, including spinal manipulation (Nielens et al., 2006; Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011) or mobilizations (Airaksinen et al., 2006; Nielens et al., 2006). Recommended treatment frequency/duration was a maximum of nine sessions over up to 12 weeks (National Institute of Health and Care Excellence, 2009).
- (4) Paracetamol or NSAIDs as therapeutic options while considering side-effects and patient preferences (Airaksinen et al., 2006; Nielens et al., 2006; Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013).
- (5) Short-term use of opioids when paracetamol or NSAIDs provided insufficient pain relief (Airaksinen et al., 2006; Nielens et al., 2006; Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013). However, it is important to take into account side-effects, risks, and patient preference (Chou et al., 2007; Livingston et al., 2009; Livingston et al., 2011; Nielens et al., 2006) and to continue only with regular re-assessments and when there is evidence of ongoing pain relief (Scottish Intercollegiate Guidelines Network, 2013).

(6) Multimodal rehabilitation that included physical and psychological interventions (e.g., cognitive/ behavioural approaches and exercise) for patients with high levels of disability or significant distress (Airaksinen et al., 2006; Nielens et al., 2006; Chou et al., 2007, 2009; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013). Recommended treatment frequency/duration was around 100 h over a maximum of up to 8 weeks (National Institute of Health and Care Excellence, 2009).

Interventions recommended by most guidelines:

- Massage (Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011; Nielens et al., 2006; Scottish Intercollegiate Guidelines Network, 2013); however, one guideline recommended against massage for chronic low back pain (Airaksinen et al., 2006). This difference is likely due to more recent evidence informing the newer guidelines' recommendations (Nielens et al., 2006; Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013).
- (2) Acupuncture (Nielens et al., 2006; Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013); however, one guideline recommended against acupuncture (Airaksinen et al., 2006). Again, this difference is likely due to more recent evidence informing the newer guidelines' recommendations (Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2009; Livingston et al., 2011; Nielens et al., 2006; Scottish Intercollegiate Guidelines Network, 2013). Recommended treatment frequency/duration was a maximum of 10 sessions over up to 12 weeks (National Institute of Health and Care Excellence. 2009).
- (3) Antidepressants as an option for pain relief, but possible side-effects (drowsiness, anticholinergic effects) should be considered (Airaksinen et al., 2006; Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011). However, one guideline recommended that antidepressants should not be used for chronic low back pain (Scottish Intercollegiate Guidelines Network, 2013), while one guideline reported conflicting

evidence on the effectiveness of antidepressants (Nielens et al., 2006).

Interventions not recommended by most guidelines:

- (1) Muscle relaxants (Chou et al., 2007; Cutforth et al., 2011: Livingston et al., 2011: Scottish Intercollegiate Guidelines Network, 2013); six guidelines made recommendations on the use of muscle relaxants (Airaksinen et al., 2006; Nielens et al., 2006; Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013). Of those, four recommended against its use (Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013) and two stated that muscle relaxants can be considered as an option for pain relief (Airaksinen et al., 2006; Nielens et al., 2006). Specifically, one guideline reported that the benefit of muscle relaxants could not be estimated due to low-quality evidence (Chou et al., 2007). Two guidelines reported that some muscle relaxants (cyclobenzaprine, benzodiazepines) may provide short-term pain relief, but cautioned against long-term use due to side-effects (drowsiness, dizziness, addiction, allergic sideeffects, reversible reduction of liver function, gastrointestinal effects) (Airaksinen et al., 2006; Nielens et al., 2006). However, evidence on the effectiveness of muscle relaxants was conflicting (Nielens et al., 2006). One guideline did not address muscle relaxants (National Institute of Health and Care Excellence, 2009).
- (2) Gabapentin (Airaksinen et al., 2006; Nielens et al., 2006; Cutforth et al., 2011; Scottish Inter-collegiate Guidelines Network, 2013); one guideline found insufficient evidence to recommend for or against gabapentin for chronic low back pain (Chou et al., 2007). Two guidelines did not address gabapentin (National Institute of Health and Care Excellence, 2009; Livingston et al., 2011). Two guidelines recommended considering gabapentin for neuropathic pain (but not chronic low back pain) (Cutforth et al., 2011; Scottish Intercollegiate Guidelines Network, 2013).
- (3) Passive modalities (Airaksinen et al., 2006; Chou et al., 2007; Cutforth et al., 2009; Cutforth et al., 2011; Nielens et al., 2006), including transcutaneous electrical nerve stimulation (TENS), laser, interferential therapy or ultrasound (Airaksinen et al., 2006; Chou et al., 2007; Cutforth et al., 2009; Cutforth et al., 2011; Nielens et al., 2006). Two guidelines found insufficient evidence for or

against laser (Chou et al., 2007; Cutforth et al., 2011) or interferential therapy (Chou et al., 2007). One guideline did not address passive modalities (Livingston et al., 2011). One guideline recommended that laser could be considered a treatment option based on inconsistent evidence (Scottish Intercollegiate Guidelines Network, 2013).

3.5 Lumbar disc herniation with radiculopathy (one high-quality guideline)

One high-quality guideline made recommendations for the noninvasive management of lumbar disc herniation with radiculopathy (Kreiner et al., 2014). Five other high-quality guidelines (Airaksinen et al., 2006; van Tulder et al., 2006; Chou et al., 2007, 2009; Livingston et al., 2011) included low back pain with leg pain in their scope, but did not have specific recommendations for the noninvasive management of lumbar disc herniation with radiculopathy. Recommended interventions:

- (1) Spinal manipulation may be an option for symptomatic relief (Kreiner et al., 2014).
- (2) A limited course of structured exercise for patients with mild to moderate symptoms. This option was based on the consensus opinion of the guideline development group (in the absence of reliable evidence; Kreiner et al., 2014).

There was insufficient evidence to make a recommendation for or against the use of traction, ultrasound, and low-level laser therapy (Kreiner et al., 2014).

4. Discussion

We conducted a systematic review of clinical practice guidelines to identify effective conservative (noninvasive) interventions for the management of acute and chronic low back pain. Most recommended interventions provide time-limited and small benefits. Based on high-quality guidelines: (1) patients with low back pain should be provided with education and encouraged to stay active and return-toactivity as tolerated; and (2) the management of acute nonspecific low back pain includes spinal manipulation (when not improving with self-care or not returning to normal activities), paracetamol or NSAIDs as indicated. Based on high-quality guidelines, the management of chronic nonspecific low back pain includes the following: (1) paracetamol or NSAIDs (although the effectiveness of paracetamol is now being challenged by new evidence); (2) shortterm use of opioids for relief of refractory, severe pain; (3) exercises; (4) manual therapy; (5) acupuncture, and (6) multimodal rehabilitation (combined physical and psychological treatment). Finally, the noninvasive management of lumbar disc herniation with radiculopathy may include spinal manipulation for symptomatic relief (Kreiner et al., 2014). Very few guidelines provided information on recommended dose and frequency of care.

Our results agree with recommended interventions identified by a previous systematic review of guidelines on low back pain (Dagenais et al., 2010). We confirmed that most passive modalities (e.g. TENS, laser, ultrasound) are not recommended for managing chronic low back pain (Dagenais et al., 2010). In addition, we found one recent high-quality guideline on lumbar disc herniation with radiculopathy published in 2012 (North American Spine Society, 2012).

However, the recommendation of paracetamol for acute low back pain is challenged by a recent highquality randomized controlled trial, which found that paracetamol did not improve recovery time compared with placebo for acute low back pain (Williams et al., 2014). Previous systematic reviews found no evidence supporting paracetamol for low back pain (Davies et al., 2008; Machado et al., 2015). Moreover, some high-quality guidelines used evidence from other conditions (e.g., osteoarthritis) to inform recommended interventions [paracetamol (Chou et al., 2007; Cutforth et al., 2011; Livingston et al., 2011) or opioids (Deyo et al., 2015)] for acute low back pain. Therefore, it is possible that using evidence for the management of other conditions, even if clinically relevant, may lead to inadequate recommendations. Given the risk of adverse events, we should reconsider the universal endorsement of paracetamol for the management of low back pain (Williams et al., 2014; Machado et al., 2015). This emphasizes that guidelines must be updated every 5 years to ensure that the most up-to-date evidence is used to inform clinical recommendations (Kung et al., 2012).

We found that high-quality guidelines lacked details about the use of acupuncture for the management of low back pain (Nielens et al., 2006; Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Cutforth et al., 2011; Livingston et al., 2011; Scottish Intercollegiate Guidelines Network, 2013). This is important because it is known that different acupuncture techniques have different levels of effectiveness (Furlan et al., 2005). Future guidelines should consider stratifying evidence by

acupuncture technique and provide clear details about the parameters for acupuncture use in patients with low back pain.

Clinical practice guidelines of low methodological quality are still being developed and published (Philippine Academy of Rehabilitation Medicine, 2011; Brosseau et al., 2012; Delitto et al., 2012). These guidelines typically fail to: (1) clearly outline selection criteria of the literature; (2) adequately describe strengths and limitations of the literature and (3) adequately describe the methods used to formulate recommendations (Ransohoff et al., 2013). Our review highlights that the next generation of high-quality guidelines must focus on applispecific populations cability to and clear implementation strategies to promote adherence. Nine of 13 eligible guidelines did not adequately address the AGREE II applicability criteria. Recent evidence suggests that favourable health and economic outcomes could be achieved if evidenceinformed decision making is used to manage low back pain (Kosloff et al., 2013). However, current clinical practice is ineffective in adhering to evidence-based guideline recommendations (Kosloff et al., 2013).

Future guidelines need to integrate the views and preferences of the target population (patients, public) into guideline development. Nine of 13 eligible guidelines did not mention whether these views and preferences were sought (Airaksinen et al., 2006; van Tulder et al., 2006; Chou et al., 2007, 2009; Cutforth et al., 2011; Livingston et al., 2011; Brosseau et al., 2012; Delitto et al., 2012; Kreiner et al., 2014). Integrating patient preferences into the guideline development process: (1) improves uptake and real-world efficiency of recommended healthcare interventions; (2) enhances consumer empowerment, and (3) informs individual patient preferences in clinical decision making (Dirksen et al., 2013; Dirksen, 2014).

The recommendations included in clinical practice guidelines typically involve the consensus of guideline expert panels who are asked to consider decision determinants, such as overall clinical benefit (effectiveness and safety), value for money (cost-effectiveness), consistency with expected societal and ethical values, and feasibility of adoption into the health system (Johnson et al., 2009). The scientific evidence serves as the foundation from which recommendations are built. Therefore, significant limitations are associated with recommendations solely developed using clinical opinions. Assembling, evaluation, and summarizing of evidence are fundamental aspects of guideline development, including a systematic review and assessment of the quality of evidence (Ransohoff et al., 2013). Recommendations based solely on opinion may be liable to biases and conflicts of interest or may not benefit patients (especially when patients' views are not considered during guideline development).

4.1 Strengths and limitations

Our review had strengths. The literature search was comprehensive, methodologically rigorous, and checked by a second librarian. We outlined detailed inclusion/exclusion criteria to identify relevant evidence-based guidelines. Pairs of independent, trained reviewers screened and critically appraised the literature. This review used a recommended critical appraisal instrument for evaluating guidelines to maintain high methodological rigour (Brouwers et al., 2010). Some guidelines lacked methodological details, and we made multiple attempts to contact authors so that our screening and critical appraisal was as accurate as possible.

The main limitation was the restriction of guidelines published in English. Most guidelines are published in the language of the target users (e.g., Haute Autorité de Santé in France or El Instituto Aragones de Ciencas de la Salud in Spain) (El Instituto Aragones de Ciencas de la Salud, 2016; Haute Autorité de Santé, 2016). It is possible that excluding guidelines published in a language other than English may have biased our results. However, it is unclear whether recommendations that are not published in English would differ from those published in English. Finally, the external validity of our results may be limited to users from English-speaking jurisdictions. A second limitation concerns the definitions used to classify acute and chronic low back pain, which varied across guidelines. Four guidelines defined chronic low back pain as pain lasting more than 3 months (Airaksinen et al., 2006; Nielens et al., 2006; van Tulder et al., 2006; Cutforth et al., 2011). Three guidelines grouped recommendations for subacute and chronic low back pain into one category (Chou et al., 2007; National Institute of Health and Care Excellence, 2009; Livingston et al., 2011). Of those, two guidelines defined subacute/chronic low back pain as pain lasting more than 4 weeks (Chou et al., 2007; Livingston et al., 2011), and one guideline defined persistent low back pain as pain lasting more than 6 weeks (National Institute of Health and Care Excellence, 2009). Finally, two guidelines did not provide a

clear definition of chronic low back pain (Chou et al., 2009; Scottish Intercollegiate Guidelines Network, 2013). The different classifications used to make recommendations for the management of low back pain complicate the evidence synthesis and may have led to the misclassification of recommendations.

5. Conclusions

Most high-quality guidelines target the noninvasive management of nonspecific low back pain and recommend education, staying active/exercise, manual therapy, and paracetamol or NSAIDs as first-line treatments. However, the endorsement of paracetamol for acute low back pain is challenged by a recent high-quality randomized controlled trial and systematic review; therefore, guidelines need updating. Some high-quality guidelines used evidence from other conditions to inform recommendations, which can lead to inadequate recommendations. Most eligible guidelines poorly addressed the applicability and implementation of recommendations. Finally, guideline developers need to involve end users during guideline development.

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Author contributions

All authors have made substantial contributions to all of the following: (1) the conception and design of the study, or acquisition of data, or analysis and interpretation of data; (2) drafting the article or revising it critically for important intellectual content; and (3) final approval of the version to be submitted.

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Supporting Information

Additional Supporting Information may be found online in the supporting information tab for this article:

Appendix S1. MEDLINE search strategy on guidelines for the management of nonspecific low back pain.

Table S1. Evidence table for high-quality guidelines.