

# Comments on a systematic review and meta-analysis of steroids for epidural injections in spinal stenosis

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## Dear editor

We read with interest the manuscript by Liu et al<sup>1</sup> a systematic review and meta-analysis of steroids for epidural injection in spinal stenosis. The results showed there was fair short- and long-term benefit for treating spinal stenosis with local anesthetic and steroids; however, the authors concluded that the meta-analysis suggested that epidural steroid injections provided limited improvement in short- and long-term benefits in lumbar spinal stenosis patients. This may be confusing to the readership. Fair evidence essentially provides moderate benefits, but the conclusion shows limited improvement.

Further, the authors included a wide variety of studies which are not applicable to the meta-analysis. Issues include studies performed with or without fluoroscopy, with short-term and long-term follow-up, with local anesthetic or without local anesthetic, inter-laminar, caudal, and transforaminal approaches, and some very small studies. Multiple studies with variable bias were also included in the meta-analysis. Further, the risk of bias assessment appears to be inappropriate. For example, both studies by Manchikanti et al,<sup>2,3</sup> even though identical, were rated differently showing variable bias. Further, multiple items were rated inaccurately which differed for each study even though they were identical. These two trials have been assessed in the past in multiple systematic reviews<sup>4-7</sup> and were rated as high quality, meeting at least 8 of 12 criteria of Cochrane review criteria which have been compressed to 7 in this assessment with one trial<sup>3</sup> scoring 4 of 7 and the second trial<sup>2</sup> scoring 3 of 7 instead of both trials scoring 6 of 7. Further, the highly rated trial by Friedly et al,<sup>8</sup> which has generated significant attention, has been met with criticism for its flawed analysis and extremely short follow-up period.<sup>9</sup>

We compliment Dr Liu et al on their effort. The analysis would be improved by better focus on appropriate inclusion criteria and precision in the establishment of criteria for homogeneity. Finally, appropriate methodologic quality or risk of bias are essential to reach unbiased clinically relevant conclusions.<sup>4-7,9,10</sup>

## Disclosure

Dr Manchikanti has provided limited consulting services to Semnur Pharmaceuticals, Incorporated, which is developing non-particulate steroids. Dr Hirsch is a consultant for Medtronic.

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## Authors' reply

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## Dear editor

Although we recognize the limitations of our meta-analysis, as noted by Dr Manchikanti, we included a wide variety of studies which may not be applicable to the meta-analysis, which are associated with issues including studies performed with or without fluoroscopy, with short-term and long-term follow-up, with local anesthetic or without local anesthetic, inter-laminar, caudal, and transforaminal approaches, and some very small studies, and some studies with variable bias were also included in the meta-analysis. We argue that we included randomized controlled trials that evaluated the efficacy and safety of epidural injections of steroids plus local anesthetic versus local anesthetic alone for the treatment of lumbar spinal stenosis (LSS) patients.<sup>1</sup> The inclusion criteria for the systematic review and meta-analysis were as follows: 1) randomized controlled trials in adults with LSS with epidural injection treatment; 2) clinical or radiological diagnosis of LSS; 3) describe neurogenic claudication with back (leg) pain and gait assessment; 4) provide the dosage and route of epidural steroid injection administration; and 5) outcomes measured, such as walking ability, pain intensity, quality of life, and global improvement. Studies evaluating radiculopathy caused by disc lesions were excluded. Studies with mixed populations were only included if the data for neurogenic claudication due to LSS were provided.<sup>1</sup>

The comments by Manchikanti and Hirsch are also important to consider. However, we wish to clarify several points. First, assessment of the methodological quality was performed independently by two investigators, the methodological quality of the trials was assessed using the Cochrane Handbook for Systematic Reviews of Interventions 5.1.0.<sup>1</sup> These two trials by Manchikanti et al<sup>2,3</sup> scoring lower scores than assessment by themselves may be caused by a hidden information source that we do not know about. Second, as noted by Manchikanti et al,<sup>4</sup> the trial by Friedly et al<sup>5</sup> which has generated significant attention in our meta-analysis has been met with criticism for its flawed analysis and extremely

short follow-up period. We also recognized some limitations of this trial such as acute pain patients being included, and that multilevel stenosis and various other factors were not identified and may have caused the risk of bias. Third, we also mentioned that in 2013, North America Spine Society's Evidence-Based Clinical Guideline Development Committee developed an evidence-based clinical guideline<sup>6</sup> for the diagnosis and treatment of degenerative LSS. They found evidence supporting the recommendation of epidural steroid injection therapy, elaborating a B recommendation in favor of it use. However, this systematic review was based on only four<sup>7-10</sup> trials. As Manchikanti and Hirsch point out, appropriate methodologic quality or risk of bias are essential to reach unbiased clinically relevant conclusions. We compliment Dr Manchikanti and Dr Hirsch on their effort. Finally, additional better and rigorous studies with long-term observation are required to elucidate the effectiveness of epidural steroid injection treatment for LSS.<sup>1</sup>

## Disclosure

The authors report no conflicts of interest in this communication.

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