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Response to the Reviewer 01436637

Little if any benefit

As I showed by citation of many articles, benefit is not little, because of analgesic effects with minimum side effects.

Animal study demonstrated harmful effect of midazolam

If this is true, all animal studies should have the histopathological damage by midazolam, but not. And there should be any clinical studies showing side effects by intrathecal or epidural midazolam, but not. As I showed in the text, morphine and lidocaine have neurotoxic in some animal studies, but used clinically. Therefore, we need further studies to conclude it.

No new message

Editorial should be in two ways; one is new material, and another is summary. This is the summary. However, this has a new message that discussion of intrathecal midazolam dose not finish yet because there are some studies to show no neurotoxicity, then we need further studies.

Spinal adjuvant

I did not say midazolam is an adjuvant. As I cited, midazolam itself has enough analgesic effect.

Possible risk of neurotoxicity

All other agents; such as morphine, fentanyl, etc. also has possible risk of neurotoxicity. However, they are used because of no neurotoxic clinical evidence. This is the same as midazolam.

I made a lot of changes according to the reviewer's suggestions. I believe that this manuscript is worth to be published in **World Journal of Anesthesiology**.

Thank you very much for reviewing our manuscript.

Tomoki Nishiyama, MD, PhD

Response to the Reviewer 01076129

Focus more on the effects of midazolam on the spinal cord

I already focused on the effects of midazolam on the spinal cord, but again rewrote most parts of the manuscript.

Toxicity of morphine and lidocaine

This is an important part to compare with the discussion of midazolam, but I shortened that part.

Neurotoxicity study in human

I changed that part.

Baclofen

I corrected the spelling.

Some words seem to have been left.

I do not understand what this means.

Nausea and vomit

I have already shown that you pointed out.

I made a lot of changes according to the reviewer's suggestions. I believe that this manuscript is worth to be published in **World Journal of Anesthesiology**.

Thank you very much for reviewing our manuscript.

Tomoki Nishiyama, MD, PhD

Response to the Reviewer 00529915

Why alternative agent?

I already showed the reason, but I changed the sentence.

Side effects

I changed the sentence.

I made a lot of changes according to the reviewer's suggestions. I believe that this manuscript is worth to be published in **World Journal of Anesthesiology**.

Thank you very much for reviewing our manuscript.

Tomoki Nishiyama, MD, PhD