

# INPLASY

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**Corresponding author:**

Zixiang Zhu

18061051006@163.com

**Author Affiliation:**

First Affiliated Hospital of Soochow University.

## The efficacy and safety of zavegepant nasal inhalation versus oral calcitonin-gene related peptide receptor antagonists in the acute treatment of migraine: A systematic review and network meta-analysis of the literature

Zhu, ZX; Tang, YB; Li, LY; Ni, HY; Liu, MR; Chen, ZQ; Wang, Z.

**ADMINISTRATIVE INFORMATION****Support** - This work was supported by Boxi Youth Natural Science Foundation (BXQN2023005) and the National Natural Science Foundation of China (No. 82171309 and No 82201445).**Review Stage at time of this submission** - Completed but not published.**Conflicts of interest** - None declared.**INPLASY registration number:** INPLASY202510064**Amendments** - This protocol was registered with the International Platform of Registered Systematic Review and Meta-Analysis Protocols (INPLASY) on 18 January 2025 and was last updated on 18 January 2025.**INTRODUCTION**

**Review question / Objective** To investigate the efficacy and safety of zavegepant nasal inhalation versus oral calcitonin-gene related peptide (CGRP) receptor antagonists for the acute treatment of migraine.

**Condition being studied** The new intranasal inhaled calcitonin gene-related peptide (CGRP) receptor antagonist, Zavegepant, has been shown to have definitive efficacy in the acute treatment of migraine. However, whether it shows superiority over other oral CGRP receptor antagonists remains to be demonstrated. Zavegepant, a new nasal inhalation calcitonin gene-related peptide (CGRP) receptor antagonist, has been proven to have a clear efficacy in the acute treatment of migraine. However, whether the efficacy of this new nasal inhalation drug is better than other oral CGRP receptor antagonists remained to be confirmed.

**METHODS**

**Search strategy** A search for eligible studies published up to 1 December 2024 was conducted on PubMed, EMBASE and the Cochrane Register of Controlled Trials (CENTRAL). PubMed, EMBASE, and The Cochrane Register of Controlled Trials (CENTRAL) were searched for eligible studies published up to December 1, 2024. The following keywords (in the title/abstract) were used: migraine, zavegepant, CGRP receptor antagonists, acute treatment of migraine.

**Participant or population** Adult participants diagnosed with migraine.

**Intervention** Zavegepant or oral CGRP receptor antagonists.

**Comparator** Placebo of the same dose.

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**Study designs to be included** RCT.

**Eligibility criteria** Exclusion criteria: comment, review, letter, animal experiment, retrospective study, case series, or case reports; no extractable data; non-English article.

**Information sources** PubMed, EMBASE, and The Cochrane Register of Controlled Trials (CENTRAL).

**Main outcome(s)** Pain freedom at 2 hours, freedom from the most bothersome symptom (MBS) at 2 hours.

**Additional outcome(s)** Pain relief at 2 hours, sustained pain freedom 2-24 hours, sustained pain freedom 2-48 hours, sustained pain relief 2-24 hours, sustained pain relief 2-48 hours, phonophobia freedom at 2 hours, photophobia freedom at 2 hours, AEs (adverse events), nausea and vomiting.

**Quality assessment / Risk of bias analysis** The risk of bias was assessed with Cochrane Collaboration tool.

**Strategy of data synthesis** Review Manager 5.3 was used. Risk ratio was analyzed with 95% confidence intervals.  $P < 0.05$  was statistically significant.

**Subgroup analysis** NA.

**Sensitivity analysis** The I<sup>2</sup> statistic was used to weigh measure heterogeneity; values less than 30% indicate "low heterogeneity," values between 30% and 50% indicate "moderate heterogeneity," and values more than 50% indicate "severe heterogeneity." Sensitivity analysis was used to investigate the stability of the consolidated data.

**Language restriction** English.

**Country(ies) involved** China.

**Keywords** migraine, zavegepant, CGRP receptor antagonists, acute treatment of migraine.

#### **Contributions of each author**

Author 1 - Zixiang Zhu.

Email: 18061051006@163.com

Author 2 - Yanbing Tang.

Email: tangyb02@163.com

Author 3 - Longyuan Li.

Email: llyuan66@163.com

Author 4 - Hanyu Ni.

Email: kirstyni@icloud.com

Author 5 - Meirong Liu.

Email: meizai26@163.com

Author 6 - Zhouqing Chen.

Email: zqchen6@163.com

Author 7 - Zhong Wang.

Email: wangzhong761@163.com