# RESEARCH

**BMC** Anesthesiology



# Comparison of anesthesia methods for intraarterial therapy of patients with acute ischemic stroke: an updated meta-analysis and systematic review



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# Abstract

**Objectives** Currently, there remains debate regarding the optimal anesthesia approach for patients undergoing intra-arterial therapy for acute ischemic stroke. Therefore, we conducted a comparative analysis to assess the effects of general anesthesia versus non general anesthesia on patient outcomes.

**Methods** The research methodology entailed comprehensive searches of prominent databases such as the Cochrane Library, PubMed, Scopus, and Web of Science, covering the period from January 1, 2010, to March 1, 2024. Data synthesis employed techniques like risk ratio or standardized mean difference, along with 95% confidence intervals. The study protocol was prospectively registered with PROSPERO (CRD42024523079).

**Results** A total of 27 trials and 12,875 patients were included in this study. The findings indicated that opting for nongeneral anesthesia significantly decreased the risk of in-hospital mortality (RR, 1.98; 95% CI: 1.50 to 2.61; p<0.00001;  $I^2 = 20\%$ ), as well as mortality within three months post-procedure (RR, 1.24; 95% CI: 1.15 to 1.34; p<0.00001;  $I^2 = 26\%$ ), while also leading to a shorter hospitalization duration (SMD, 0.24; 95% CI: 0.15 to 0.33; p<0.00001;  $I^2 = 44\%$ ).

**Conclusion** Ischemic stroke patients who undergo intra-arterial treatment without general anesthesia have a lower risk of postoperative adverse events and less short-term neurological damage. In routine and non-emergency situations, non-general anesthetic options may be more suitable for intra-arterial treatment, offering greater benefits to patients. In addition to this, the neuroprotective effects of anesthetic drugs should be considered more preoperatively and postoperatively.

Keywords Stroke, Intra-arterial therapy, General anesthesia, Local anesthesia, Conscious sedation

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#### Introduction

Stroke is a globally prevalent disease marked by high mortality and disability rates. It is classified into two main types based on its pathological features: ischemic stroke and hemorrhagic stroke. Ischemic stroke is the most common type of stroke, accounting for approximately 70% of all strokes [1]. The key to the treatment of acute ischemic stroke is to open the blocked blood vessels as early as possible and save the ischemic penumbra. The treatment method for early vascular recanalization of acute ischemic stroke that has been used for a long time in the past is mainly intravenous recombinant tissue plasminogen activator (rt-PA) thrombolysis [2–4]. Some studies have shown that intravenous rt-PA thrombolysis within 4.5 h of onset has clear benefits, and the earlier the thrombolysis, the greater the benefit [5]. However, intravenous thrombolysis has a strict time window limit, and the number of patients who can benefit from it is less than 3% of ischemic stroke patients. At the same time, there is still huge room for optimization of its therapeutic effect. Therefore, scholars around the world are exploring the intra-arterial therapy (IAT) of ischemic stroke [6]. Studies have shown that IAT based on mechanical thrombectomy can bring clear benefits and has now become the standard treatment for acute ischemic stroke in addition to intravenous thrombolysis [7-9].

As technology and materials improve, new problems arise. So far, the optimal anesthesia regimen for IAT in acute ischemic stroke remains controversial. General anesthesia (GA), as a widely-used method, offers several benefits. It effectively immobilizes the patient, minimizing involuntary movements. Additionally, it mitigates the risk of aspiration by managing the airway effectively, and it allows for superior control over circulation. Local anesthesia (LA) is also widely used in neurology-related surgeries. With the introduction of the concept of comfortable medicine, simple LA at the puncture point is no longer used, and is replaced by conscious sedation (CS) and monitored anesthesia care (MAC). CS has the characteristics of rapid onset of action and short preoperative preparation time, while MAC can better monitor hemodynamics and other vital signs of the patient to ensure safety. A retrospective study found that the use of GA during intra-arterial therapy had a more pronounced adverse effect on clinical outcomes than CS [10]. The results of several randomized controlled trials (RCTs) show that there is no significant difference in clinical outcomes between CS or GA for intra-arterial therapy [11–13]. Another meta-analysis showed that patients with anterior circulation stroke treated under GA may have better clinical neurological outcomes [14]. There is still controversy over which anesthesia regimen is best for intra-arterial therapy. Therefore, we collected relevant articles in recent years to conduct an updated meta-analysis and provide new guidance for clinical practice.

#### Methods

The research adhered to the guidelines outlined in the Preferred Reporting Items for Systematic Reviews and Meta-Analyses Protocols (PRISMA) statement [15]. Furthermore, the protocol has been officially registered in the International Prospective Systematic Reviews Registry database with the registration number CRD42024523079.

#### Sources of data and search strategy

A comprehensive search was carried out across several databases including PubMed, Web of Science, Scopus, and the Cochrane Library, spanning from January 2010 to March 2024, to identify studies related to acute ischemic stroke and anesthesia. Part of the search strategy is as follows: [(Stroke) OR (Cerebrovascular Accident)] AND [(Anesthesia, General) OR (Monitored anesthesia care) OR (Conscious Sedation)] in title/abstract. Furthermore, citations from articles were extracted to pinpoint relevant studies that might not have been initially captured during the literature search. The detailed search strategy is outlined in a Word document included within the supplementary materials.

#### Inclusion and exclusion criteria

The inclusion criteria were established in accordance with the PICOS approach. These criteria include: (1) Original clinical studies contrasting GA with non-GA; (2) Participants aged over 18 years undergoing intraarterial therapy, with baseline data and comorbidities not significantly special or high-risk; (3) Inclusion of pertinent clinical outcomes essential for this investigation. The exclusion criteria are as follows: (1) Literature types not classified as clinical trials, such as reviews, letters, and conference abstracts; (2) Studies lacking a comparison between GA and non-GA in intra-arterial therapy; (3) Insufficient data or inability to transform data into a usable format. Two authors independently reviewed and selected studies based on these predetermined criteria. Any discrepancies were resolved through discussion with a third party.

#### Data collection and quality assessment

Two reviewers autonomously performed data extraction. Any disparities were resolved through consensus or by seeking input from a third party. The extracted data encompassed various details, including the primary author, publication year, sample size, participant demographics (age, gender), comorbidities, National Institutes of Health Stroke Scaleas (NIHSS) score, as well as primary and secondary outcomes. In accordance with the "Randomized Trial Bias Risk Assessment Tool" as outlined in the Cochrane Handbook, the quality assessment of the randomized controlled trials (RCTs) encompasses several domains. These encompass allocation concealment, randomization method, blinding procedures for both investigators and participants, blinding of outcome assessors, selective reporting, completeness of data, and identification of other potential biases. The overall risk of bias assessment can lead to categorizations of low, unclear, or high risk of bias [16].

For retrospective studies, quality assessment was carried out utilizing the Newcastle-Ottawa Scale (NOS) by two independent reviewers. The assessment entailed evaluating three key aspects: selection bias, comparability, and exposure. Each aspect featured specific evaluation criteria, with stars allocated accordingly. The highest score attainable for comparability is two stars.

## **Outcomes and definitions**

The primary outcomes included functional independence at 3 months, in-hospital mortality, and mortality at 3 months. Functional independence at 3 months is defined as achieving an modified Rankin Scale (mRS) score of 0 to 2. Secondary outcomes were successful reperfusion, intracranial hemorrhage, pneumonia, NIHSS score after 24 h, vascular perforation, progressive ischemic stroke, and length of hospital stay. Successful reperfusion was defined as an modified Thrombolysis in Cerebral Infarction (mTICI) score of 2b or 3 indicating reperfusion of more than 50% of the affected area.

# Statistical analysis

All data underwent analysis using Review Manager (Rev-Man) version 5.4 (The Cochrane Collaboration, Copenhagen, Denmark) and Stata SE 16.0 (Stata Corporation, College Station, TX, USA). For dichotomous data, risk ratios (RR) with corresponding 95% confidence intervals (CI) were computed, while for continuous data, standard mean differences (SMD) with 95% CI were estimated. Both fixed and random effects models were employed to accommodate methodological and clinical heterogeneity. Heterogeneity among studies was evaluated using the Q-test and I<sup>2</sup> statistic, with significant heterogeneity defined as p < 0.1 or I<sup>2</sup>>50%. Subgroup and metaregression analyses were conducted to explore potential sources of heterogeneity. Publication bias was assessed through funnel plots, with Egger's test employed when at least 10 studies were included. TSA 0.9.5.10 beta software was used to conduct trial sequential analysis (TSA) of clinical efficacy to reduce the occurrence of random errors, determine the reliability of the conclusions, and estimate the sample size required for meta-analysis. A significance level of  $\alpha = 0.05$  was utilized for all analyses.

Sensitivity analysis was conducted to assess result robustness and to identify potential sources of heterogeneity.

#### Results

# Literature selection

A total of 1219 pieces of literature were identified across various databases. Following the removal of 114 duplicate studies, a preliminary screening excluded 1105 studies. Subsequently, 41 articles underwent full-text evaluation, ultimately resulting in the inclusion of 27 trials for final analysis. Within this selection, three articles were omitted as they did not constitute original clinical studies, seven articles were disregarded due to the absence of a comparison between general anesthesia (GA) and non-GA, and five articles were excluded either due to the lack of relevant results or the inability to convert the data into a usable format. The specific screening process is detailed in Fig. 1. Among the 27 included articles, 12 were RCTs and 15 were cohort studies [11, 12, 17–41].

#### Baseline characteristic and quality assessment

The trials analyzed in this study were all published post-2010, featuring sample sizes ranging from 40 to 4429 individuals. In total, 12,875 participants were included, with an average age of 69.5 years. The baseline demographic characteristics and comorbidities of the patients are detailed in Table 1.

After a thorough quality assessment of twelve randomized controlled trials, two were determined to have a high risk of bias, five to have a low risk of bias, and the remaining five were considered to have an unclear risk of bias (Figure S1). After assessing the quality of the remaining fifteen retrospective studies, we found that all studies had above-average NOS scores. Each study scored more than five stars and met the criteria for inclusion in the metaanalysis. The conclusive results are detailed in Table S1.

#### Main outcomes

Non-GA is associated with a smaller risk of in-hospital death (RR, 1.98; 95% CI: 1.50 to 2.61; p < 0.00001;  $I^2 = 20\%$ ) and three-month mortality (RR, 1.24; 95% CI: 1.15 to 1.34; p<0.00001; I<sup>2</sup>=26%) than GA. Non-GA is associated with higher successful reperfusion (RR, 1.06; 95% CI: 1.01 to 1.11; p=0.02;  $I^2=60\%$ ) and lower risk of progressive ischemic stroke (RR, 1.41; 95% CI: 1.10 to 1.79 ; p=0.006; I<sup>2</sup>=27%). In addition, patients without GA had lower NIHSS scores 24 h after surgery (SMD, 0.13; 95% CI: 0.01 to 0.25; *p*=0.03; I<sup>2</sup>=54%), but the mRS score results at three months (RR, 0.89; 95% CI: 0.79 to 0.99 ; p=0.04; I<sup>2</sup>=55%) showed that patients with GA had better outcomes. Patients without GA had shorter hospital stays (SMD, 0.24; 95% CI: 0.15 to 0.33; p<0.00001;  $I^2$ =44%).There may be a lower risk of vascular perforation and a higher risk of intracranial hemorrhage after



Fig. 1 Preferred reporting items for systematic reviews and meta-analyses (PRISMA) flowchart of selection

general anesthesia, but this result is not statistically significant, and there is no significant difference in the risk of postoperative pneumonia. Relevant results are summarized in Tables 2 and 3, and forest plots are shown in the supplementary material.

## Subgroups analysis

We conducted subgroup analysis on mRS $\leq 2$  after three months, three-month mortality, in-hospital death, successful reperfusion and length of hospital stay according

to different study types. The trends reported in RCT studies and non-RCT studies are basically the same, and no significant difference. In addition, we found that different study types may be one of the sources of heterogeneity in three-month mortality, in-hospital death, successful reperfusion and length of hospital stay.

Considering the relationship between surgical volume and outcomes for complex and high-risk surgeries, differences in sample size may have influenced the results. Therefore, we grouped mRS $\leq 2$  after three months,

Reference	Sam- ple size	Age	Male	HT	AF	DM	HL	Smoking	CAD	HF	NIHSS
Abou [17]	281	67.2±15.0	145 (51.6)	211 (75.1)	112 (39.9)	72 (25.6)	138 (49.1)	87 (31.0)	85 (30.2)	NA	18.1±6.6
Bekelis [18]	1174	67.3±15.0	559 (47.6)	791 (67.4)	NA	289 (24.6)	475 (40.5)	151 (12.9)	322 (27.4)	313 (26.7)	NA
Berkhemer [19]	216	$65.0 \pm 16.4$	126 (58.3)	NA	62 (28.7)	31 (14.4)	NA	NA	NA	NA	$17.6 \pm 4.9$
Cappellari [20]	4429	71.1±14.0	2241 (50.6)	2442 (55.1)	1155 (26.1)	641 (14.5)	511 (11.5)	785 (17.7)	382 (8.6)	267 (6.0)	17.6±5.4
Chabanne [21]	273	71.6±13.8	131 (48.0)	167 (61.2)	NA	38 (13.9)	NA	NA	NA	29 (10.6)	15.5±6.7
Farag [22]	358	67.7±15.0	174 (48.6)	323 (90.2)	NA	133 (37.2)	86 (24.0)	NA	NA	103 (28.8)	15.2±7.3
Goldhoorn [23]	1376	$69.5\pm15.0$	742 (53.9)	696 (50.6)	308 (22.3)	231 (16.8)	414 (30.1)	313 (22.7)	NA	NA	$15.6 \pm 6.4$
Hu [24]	139	$72.0 \pm 7.1$	72 (51.8)	65 (46.8)	51 (36.7)	21 (15.1)	49 (35.3)	40 (28.8)	NA	NA	NA
Jagani [25]	99	66.1±12.4	52 (52.5)	75 (75.8)	31 (31.3)	17 (17.2)	NA	46 (46.5)	27 (27.3)	NA	NA
Janssen [26]	84	$69.8 \pm 12.5$	38 (45.2)	64 (76.2)	42 (50.0)	10 (11.9)	22 (26.2)	21 (25.0)	NA	NA	NA
John [27]	190	$67.0 \pm 15.2$	83 (43.7)	137 (72.1)	75 (39.5)	45 (23.7)	85 (44.7)	NA	NA	NA	$15.8 \pm 6.6$
Just [28]	109	61.9	67 (35.3)	61 (56.0)	NA	19 (17.4)	NA	52 (47.7)	NA	NA	13.1
Li [29]	109	66.1±16.3	53 (48.6)	79 (72.5)	32 (29.4)	27 (24.8)	51 (46.8)	NA	38 (34.9)	NA	16.0±6.3
Li [30]	636	NA	359 (56.4)	380 (59.7)	294 (46.2)	120 (18.9)	NA	NA	NA	NA	NA
Liang [31]	87	62.0±12.0	71 (81.6)	63 (72.4)	14 (16.1)	23 (26.4)	31 (35.6)	53 (60.9)	13 (14.9)	NA	15.7±5.7
Maurice [32]	351	$71.7 \pm 12.6$	194 (55.3)	221 (63.0)	107 (30.5)	49 (14.0)	NA	NA	NA	NA	$16.0\pm5.5$
Mundiyanapurath [33]	44	72.3±14.1	19 (43.2)	NA	NA	NA	NA	NA	NA	NA	19.2±7.1
Peng [34]	149	$63.5 \pm 12.9$	92 (61.7)	84 (56.4)	60 (40.3)	14 (9.4)	7 (4.7)	43 (28.9)	NA	NA	$16.0\pm5.9$
Pop [35]	361	$73.0 \pm 15.1$	169 (46.8)	242 (67.0)	NA	70 (19.4)	123 (34.1)	57 (15.8)	NA	NA	$15.2 \pm 7.2$
Ren [36]	90	$69.2 \pm 6.1$	50 (55.6)	37 (41.1)	9 (10.0)	11 (12.2)	6 (6.7)	NA	NA	NA	$13.6\pm3.8$
Schonenberger 2016	150	71.5±13.8	90 (60.0)	107 (71.3)	72 (48.0)	34 (22.7)	44 (29.3)	22 (14.7)	NA	38 (25.3)	17.0±3.8
Simonsen [12]	128	71.4±11.4	66 (51.6)	71 (55.5)	51 (39.8)	18 (14.1)	NA	40 (31.3)	NA	NA	$17.5 \pm 5.4$
Sun [38]	40	$63.2 \pm 19.2$	26 (65.0)	17 (42.5)	12 (30.0)	9 (22.5)	NA	NA	7 (17.5)	NA	$13.7\pm6.0$
Vandenberg 2015	348	$61.3 \pm 14.7$	184 (52.9)	167 (48.0)	90 (25.9)	49 (14.1)	85 (24.4)	NA	NA	NA	NA
Wagner [ <mark>39</mark> ]	1284	$71.6 \pm 13.6$	667 (51.9)	895 (69.7)	512 (39.9)	222 (17.3)	819 (63.8)	270 (21.0)	NA	NA	$14.0\pm7.7$
Wu [41]	187	$64.1\pm10.9$	127 (67.9)	97 (51.9)	53 (28.3)	46 (24.6)	39 (20.9)	79 (42.2)	NA	NA	$14.3\pm6.6$
Wu [40]	183	59.7 + 11.6	148 (80.9)	141 (77.0)	31 (16.9)	54 (29.5)	31 (16.9)	87 (47.5)	NA	NA	22.0 + 14.5

#### Table 1 Basic information included in the studies

HT, Hypertension; AF, Atrial fibrillation; DM, Diabetes mellitus; HL, Hyperlipidemia; CAD, Coronary artery disease; HF, heart failure; NIHSS, National Institutes of Health Stroke Scale; NA, not applicable

Data are expressed as mean ± SD or mean or n (%)

three-months mortality and recanalization success according to different sample size levels and conducted subgroup analysis. The results show that differences in sample size do not significantly affect the results and that sample size is not a source of heterogeneity (Figs. 2, 3 and **4**).

# **Meta-regression**

A random effects multivariable meta-regression analysis was performed to examine the association between mRS $\leq$ 2 three months post-stroke, NIHSS score at 24 h and successful reperfusion. Factors including age, gender, hypertension, hyperlipidemia, smoking, and NIHSS score were taken into account. None of these factors were identified as potential sources of heterogeneity. Detailed results are presented in the supplementary material. Considering the possible influence of sample size on the results, we also conducted a meta-regression with sample size as the covariate for mRS  $\leq 2$  after three months, three-months mortality and recanalization success. The results showed that sample size was not the source of heterogeneity.

Outcomes	No. of studies	Participants	Statistical Method	Effect Estimate	l <sup>2</sup>	<i>p</i> -value
mRS ≤ 2 after th	ree months					
RCT	8	2927	RR (M-H, Random)	0.94 [0.79, 1.11]	53%	0.45
Non-RCT	10	6378	RR (M-H, Random)	0.84 [0.71, 1.00]	58%	0.05
Total	18	9305	RR (M-H, Random)	0.89 [0.79, 0.99]	55%	0.04
Three-months r	nortality					
RCT	9	2644	RR (M-H, Fixed)	1.02 [0.87, 1.18]	0	0.84
Non-RCT	10	8288	RR (M-H, Fixed)	1.32 [1.21, 1.44]	24%	< 0.00001
Total	18	10,932	RR (M-H, Fixed)	1.24 [1.15, 1.34]	26%	< 0.00001
In-hospital deat	h					
RCT	3	327	RR (M-H, Fixed)	1.04 [0.53, 2.04]	0	0.90
Non-RCT	4	766	RR (M-H, Fixed)	2.33 [1.71, 3.19]	0	< 0.00001
Total	7	1093	RR (M-H, Fixed)	1.98 [1.50, 2.61]	20%	< 0.00001
Successful repe	rfusion					
RCT	11	2164	RR (M-H, Fixed)	1.07 [1.02, 1.12]	36%	0.007
Non-RCT	10	6393	RR (M-H, Fixed)	1.01 [0.98, 1.04]	68%	0.36
Total	21	8557	RR (M-H, Random)	1.06 [1.01, 1.11]	60%	0.02
NIHSS score afte	er 24 h					
Total	9	3319	SMD (IV, Random)	0.13 [0.01, 0.25]	54%	0.03
(All RCT)						
Progressive isch	emic stroke					
RCT	3	1865	RR (M-H, Random)	1.40 [1.06, 1.86]	67%	0.02
Non-RCT	3	718	RR (M-H, Fixed)	1.42 [0.88, 2.29]	0	0.16
Total	6	2583	RR (M-H, Fixed)	1.41 [1.10, 1.79]	27%	0.006
Length of hospi	ital stay					
RCT	5	2012	SMD (IV, Fixed)	0.10 [-0.05, 0.25]	19%	0.21
Non-RCT	2	699	SMD (IV, Fixed)	0.32 [0.21, 0.43]	0	< 0.00001
Total	7	2012	SMD (IV, Fixed)	0.24 [0.15, 0.33]	44%	< 0.00001

|--|

 Table 3
 Other outcomes and statistical results

Outcomes	Studies	Participants	Risk ratio	95% CI	<i>p</i> -value	Corresponding figure
Intracerebral hemorrhage	18	10,710	1.08	0.93-1.26	0.32	Figure <mark>S9</mark>
Pneumonia	11	3058	1.01	0.85-1.20	0.93	Figure S10
Vessel perforations	6	1299	0.51	0.24-1.08	0.08	Figure S11

#### Publication bias and sensitivity analysis

The funnel plots for all findings revealed no substantial evidence of publication bias. Furthermore, both Egger's and Begg's tests were conducted, confirming the absence of publication bias across all outcomes. For outcomes with fewer than 10 included studies, publication bias was not examined. Detailed funnel plots and test outcomes are available in the supplementary materials. Additionally, the sensitivity analysis underscores the robustness of our results (accessible in the supplementary materials).

# TSA

This study conducted TSA for mRS $\leq 2$  after three months and three-months mortality, setting the type I error rate  $\alpha = 0.05$ , the information axis as the cumulative sample size, the statistical power of 80%, and the sample size as the required information size (RIS), see Figure S15 and Figure S16. As a result, the Z-curve crossed both the traditional boundary and the TSA boundary,

and its cumulative information volume reached RIS. It shows that under the effect of RR=0.89, non-GA has clear evidence for improving the three-month neurological prognosis of patients. With the effect of RR=1.24, the evidence that GA can improve the three-month mortality rate of patients is conclusive.

# Discussion

The most effective anesthetic approach for IAT in ischemic stroke continues to be a subject of debate and contention among medical professionals. Current guidelines advise tailoring decisions to individual patient characteristics, yet they do not offer precise recommendations [3]. Previous studies have yielded varying conclusions regarding the advantages and disadvantages of different anesthesia methods for ischemic stroke patients undergoing IAT. Möhlenbruch et al [42]:s study, comprising 111 patients who received IAT for posterior circulation stroke, found that patients under CS exhibited

	GA		Non-0	GA		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight I	M-H, Random, 95% CI	M-H, Random, 95% CI
3.1.1 sample size < 20	0						2 1 1 2 2 1 1 2 1 2 1 2 1 2 1 2 1 2 1 2
Jagani 2015	6	38	22	61	1.7%	0.44 [0.20, 0.98]	
John 2014	10	91	18	99	2.0%	0.60 [0.29, 1.24]	
Just 2016	14	42	31	67	3.7%	0.72 [0.44, 1.19]	
Liang 2022	21	43	24	44	4.8%	0.90 [0.60, 1.35]	
Peng 2019	27	44	56	105	7.0%	1.15 [0.86, 1.55]	
Schonenberger 2016	27	73	14	77	3.1%	2.03 [1.16, 3.56]	
Sun 2018	11	20	10	20	2.8%	1.10 [0.61, 1.99]	
Wu 2019	40	75	56	112	7.3%	1.07 [0.81, 1.41]	-
Wu 2020	36	112	25	71	4.7%	0.91 [0.60, 1.38]	
Subtotal (95% CI)		538		656	37.2%	0.97 [0.78, 1.20]	•
Total events	192		256				
Heterogeneity: Tau <sup>2</sup> = 0	.05; Chi?:	= 15.52	, df = 8 (F	P = 0.05	5); I <sup>2</sup> = 48%	G	
Test for overall effect: Z	= 0.28 (P	= 0.78)	)				
3.1.2 200 <sample size<="" td=""><td>e&lt;1000</td><td></td><td></td><td></td><td></td><td></td><td></td></sample>	e<1000						
Abou 2014	70	196	45	85	7.5%	0.67 (0.51, 0.89)	
Berkhemer 2016	18	79	52	137	4.1%	0.60 [0.38, 0.95]	
Chabanne 2023	45	135	54	138	6.5%	0.85 10.62, 1,171	
Farag 2022	16	104	47	254	3.5%	0.83 [0.49, 1.40]	
LI 2023	68	207	166	429	8.7%	0.85 [0.68, 1.07]	
Pop 2021	50	97	34	97	6.2%	1.47 [1.05, 2.05]	
Vandenberg 2015	10	70	72	278	2.7%	0.55 [0.30, 1.01]	
Subtotal (95% CI)		888		1418	39.3%	0.82 [0.65, 1.03]	•
Total events	277		470				22
Heterogeneity: Tau <sup>2</sup> = 0	.06; Chi <sup>a</sup> :	= 17.29	df = 6 (f	<sup>&gt;</sup> = 0.00	08); I <sup>P</sup> = 659	%	
Test for overall effect Z	= 1.71 (P	= 0.09)	)				
3.1.3 sample size>10	00						
Cappellari 2020	803	2013	1131	2416	13.1%	0.85 [0.80, 0.91]	•
Goldhoorn 2020	117	381	358	995	10.4%	0.85 [0.72, 1.01]	
Subtotal (95% CI)		2394	150763	3411	23.5%	0.85 [0.80, 0.91]	•
Total events	920		1489				
Heterogeneity, Tau <sup>2</sup> = 0	.00: Chi#:	= 0.00.	df = 1 (P	= 0.99)	: I <sup>2</sup> = 0%		
Test for overall effect Z	= 4.93 (P	< 0.00	001)				
Total (95% CI)		3820		5485	100.0%	0.89 [0.79, 0.99]	•
Total events	1399	0000	2215	0 100	1001010	eres fair al aigal	
Heterogeneity Tau <sup>2</sup> = 0	02 Chil	- 37 40	df=17	P = 0.0	103) P = 54	596	
Test for overall effect 7	= 2.08 /P	= 0.04	, ui - 17	v - 0.0	, - Ju		0.1 0.2 0.5 1 2 5 10
Test for subarous differ	= 2.00 (F	-0.04	18 df = 2	P = 0	49) P = 0%		GA Non-GA
restion suburous unler	entres: U	- 1.3	0.01 = 2	$u^{*} = 0.$	-0.1 - 0.8		

Fig. 2 Subgroup analysis of mRS ≤ 2 after three months based on different sample size

significantly lower mRS scores 90 days post-treatment compared to those under GA. Conversely, two other studies indicated that GA was associated with poorer functional outcomes at the three-month mark [43, 44]. However, some research suggests that anesthesia methods may not significantly correlate with clinical functional outcomes. For instance, Nogueira et al [45].'s case-control study involving 215 patients with posterior circulation stroke who underwent IAT revealed similar rates of successful reperfusion, functional independence, hemorrhagic transformation, and mRS scores between the GA and CS groups. Similarly, Peng et al [46].'s study, encompassing 639 patients with basilar artery occlusion undergoing IAT, found no statistically significant differences in favorable functional outcomes, mortality, hemorrhagic transformation, or three-month mRS scores among patients undergoing GA, LA or CS.

Our study encompassed a substantial volume of documents, with primary findings indicating notable benefits associated with non-GA compared to GA Notably, the GA cohort exhibited elevated risks of mortality, disease advancement, and prolonged hospitalization. Intriguingly, while the GA group demonstrated improved mRS scores at three months post-surgery compared to the non-GA group, their NIHSS scores 24 h after surgery were inferior. There was acceptable heterogeneity in some of our outcomes, and we also conducted subgroup analysis, meta-regression, and sensitivity analysis to explore heterogeneity. The results indicate that different study types may be one of the sources of heterogeneity in three-month mortality, in-hospital death, successful

	GA		Non-(	5A		<b>Risk Ratio</b>	Ris	k Ratio	
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% CI	M-H, Fi	xed, 95% CI	
3.2.1 sample size < 200	)								
Jagani 2015	12	38	14	61	1.1%	1.38 [0.71, 2.65]			
Just 2016	18	42	14	67	1.1%	2.05 [1.15, 3.67]			
Liang 2022	10	43	9	44	0.9%	1.14 [0.51, 2.52]	-	+	
Peng 2019	5	44	13	105	0.8%	0.92 [0.35, 2.42]			
Ren 2020	9	48	9	42	1.0%	0.88 [0.38, 2.00]			
Schonenberger 2016	18	73	19	77	1.9%	1.00 [0.57, 1.75]	-	+-	
Simonsen 2018	5	65	8	63	0.8%	0.61 [0.21, 1.75]		+-	
Sun 2018	1	20	6	20	0.6%	0.17 [0.02, 1.26]		+	
Wu 2019	14	75	17	112	1.4%	1.23 [0.65, 2.34]	11		
Wu 2020	34	112	21	71	2.6%	1.03 [0.65, 1.62]		+	
Subtotal (95% CI)		560		662	12.3%	1.09 [0.88, 1.35]		•	
Total events	126		130						
Heterogeneity: Chi <sup>2</sup> = 10	).21, df=	9 (P = 1	0.33); I <sup>#</sup> =	12%					
Test for overall effect: Z :	= 0.75 (P	= 0.45	)						
3.2.2 200 <sample size<="" td=""><td>&lt;1000</td><td></td><td></td><td></td><td></td><td></td><td></td><td></td><td></td></sample>	<1000								
Abou 2014	67	196	20	85	2.9%	1.45 [0.95, 2.23]			
Chabanne 2023	25	135	23	138	2.3%	1.11 [0.66, 1.86]		<u>+-</u>	
Maurice 2022	31	174	28	177	2.9%	1.13 [0.71, 1.80]		+-	
Pop 2021	21	97	25	97	2.6%	0.84 [0.51, 1.40]		+	
Vandenberg 2015	15	70	46	278	1.9%	1.30 [0.77, 2.18]		+	
Subtotal (95% CI)		672		775	12.5%	1.17 [0.94, 1.45]		•	
Total events	159		142						
Heterogeneity: Chi# = 2.4	82, df = 4	(P = 0.	59); P = (	9%					
Test for overall effect Z :	= 1.39 (P	= 0.16	)						
3.2.3 sample size>100	00								
Bekelis 2017	125	441	147	733	11.4%	1.41 [1.15, 1.74]		-	
Cappellari 2020	406	2013	398	2416	37.2%	1.22 [1.08, 1.39]		-	
Goldhoorn 2020	107	381	267	995	15.2%	1.05 (0.86, 1.27)		+	
Wagner 2022	264	851	83	433	11.3%	1.62 [1.30, 2.01]		-	
Subtotal (95% CI)		3686	ł	4577	75.1%	1.28 [1.17, 1.39]		+	
Total events	902		895					57	
Heterogeneity: Chi# = 10	).08. df=	3 (P = 1	0.02); 1=	70%					
Test for overall effect: Z :	= 5.58 (P	< 0.00	001)						
Total (95% CI)		4918		6014	100.0%	1.24 [1.15, 1.34]		•	
Total events	1187		1167		Chelosoper 2				
Heterogeneity: Chi <sup>2</sup> = 24	1.33. df=	18 (P =	0.14): 17	= 26%			terre	+ +	
Test for overall effect 7:	= 5.62 (P	< 0.00	001)				0.01 0.1	1 10	100
Test for subgroup differe	ences: Cl	ni <sup>2</sup> = 2.1	19. df = 2	(P = 0)	33),   <sup>2</sup> = 8	.8%	G	A NON-GA	

Fig. 3 Subgroup analysis of three-months mortality based on different sample size

reperfusion and length of hospital stay. In addition, when excluding the study by Wagner et al [39], the I<sup>2</sup> for threemonth mortality dropped to 4%, p<0.0001, and the RR was 1.19. When the study by Farag et al [22] is eliminated, the I<sup>2</sup> of in-hospital death drops to 0, p=0.001, and the RR is 1.69. Clinical heterogeneity caused by different treatment plans, anesthesia plans, and nursing plans adopted by different centers is also one of the sources of heterogeneity in this study. Overall, our heterogeneity is small and acceptable and does not affect the reliability of the results of this study.

Patients who do not receive GA may experience quicker neurological recovery within 24 h post-surgery, as indicated by lower NIHSS scores. Conversely, patients undergoing surgery with GA may enjoy more consistent and enduring treatment outcomes over the course of the three-month observation period, potentially resulting in improved performance on mRS scores. In the immediate aftermath of surgery, patients who did not receive GA demonstrated superior performance on NIHSS scores, potentially due to a prompt restoration of neurological function post-operation. Conversely, patients who underwent surgery under GA exhibited improved mRS scores several months later, suggesting they may have benefited from the neuroprotective properties of GA over an extended duration, facilitating a more favorable recovery trajectory. Such distinctions could arise from varying physiological and neurological responses at different

<tbody<tr>Study or SubgroupE3.3.1 sample size&lt;200Hu 2021Janssen 2016John 2014Liang 2022Mundiyanapurath 2015Peng 2019Ren 2020Schonenberger 2016Simonsen 2018Wu 2019Wu 2020Subtotal (95% CI)Total eventsHaterapealie Chiller 15 12</tbody<tr>	GA		Non-0	βA		Risk Ratio	Risk Ratio
3.3.1 sample size <200 Hu 2021 Janssen 2016 John 2014 Liang 2022 Mundiyanapurath 2015 Peng 2019 Ren 2020 Schonenberger 2016 Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	Events	Total	Events	Total	Weight	M-H, Fixed, 95% CI	M-H, Fixed, 95% CI
Hu 2021 Janssen 2016 John 2014 Liang 2022 Mundiyanapurath 2015 Peng 2019 Ren 2020 Schonenberger 2016 Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events						- Carlos Constanting	
Janssen 2016 John 2014 Liang 2022 Mundiyanapurath 2015 Peng 2019 Ren 2020 Schonenberger 2016 Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	53	72	51	67	1.8%	0.97 [0.80, 1.17]	
John 2014 Liang 2022 Mundiyanapurath 2015 Peng 2019 Ren 2020 Schonenberger 2016 Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	43	53	25	31	1.0%	1.01 [0.81, 1.25]	
Liang 2022 Mundiyanapurath 2015 Peng 2019 Ren 2020 Schonenberger 2016 Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	52	91	47	99	1.5%	1,20 [0.92, 1,58]	
Mundiyanapurath 2015 Peng 2019 Ren 2020 Schonenberger 2016 Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	41	43	34	44	1.1%	1.23 [1.04, 1.47]	
Peng 2019 Ren 2020 Schonenberger 2016 Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	20	29	8	15	0.3%	1,29 (0,76, 2,20)	
Ren 2020 Schonenberger 2016 Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	37	44	89	105	1.7%	0.99 (0.85, 1.15)	
Schonenberger 2016 Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	42	48	36	42	1.3%	1.02 (0.87, 1.20)	
Simonsen 2018 Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	65	73	62	77	2.0%	1.11 [0.97, 1.27]	
Sun 2018 Wu 2019 Wu 2020 Subtotal (95% CI) Total events	50	65	38	63	1.3%	1.28 [1.00, 1.62]	
Wu 2019 Wu 2020 Subtotal (95% CI) Total events	19	20	13	20	0.4%	1.46 [1.04, 2.05]	
Wu 2020 Subtotal (95% CI) Total events	62	75	91	112	2.4%	1.02 [0.89, 1.17]	
Subtotal (95% CI) Total events	98	112	63	71	2.6%	0.99 (0.88, 1.10)	
Total events		725		746	17.5%	1.08 [1.02, 1.14]	•
Hotorononoity Chil- 15 13	582		557				
$\square$ B B B I U G B I B I B I B I B I B I B I B I B I B	df = 1	1 (P = 0)	(18): P=	27%			
Test for overall effect $7 = 2$	76 (P =	0.006					
		,					
3.3.2 200 <sample size<1<="" td=""><td>1000</td><td></td><td></td><td></td><td></td><td></td><td></td></sample>	1000						
Abou 2014	144	196	62	85	2.9%	1.01 (0.86, 1.18)	
Berkhemer 2016	41	79	86	137	2.1%	0.83 [0.64, 1.06]	
Chabanne 2023	115	135	107	138	3.5%	1.10 [0.98, 1.23]	+
Farag 2022	72	104	205	254	3.9%	0.86 [0.74, 0.99]	
LI 2023	169	207	340	429	7.3%	1.03 [0.95, 1.12]	+
Maurice 2022	144	174	131	177	4.3%	1.12 [1.00, 1.25]	
Pop 2021	93	97	68	97	2.3%	1.37 [1.19, 1.57]	
Vandenberg 2015	34	70	113	278	1.5%	1.19 (0.90, 1.58)	
Subtotal (95% CI)		1062		1595	27.8%	1.05 [1.00, 1.10]	•
Total events	812		1112				2 C C
Heterogeneity Chi <sup>2</sup> = 29.16	df = 7	(P = 0.1)	0001): (*:	76%			
Test for overall effect: Z = 1.	88 (P =	0.06)					
3.3.3 sample size>1000			1	- de la bla	-		1
Cappellari 2020	1509	2013	1817	2416	54.7%	1.00 [0.96, 1.03]	7
Subtotal (95% CI)		2013		2416	54.7%	1.00 [0.96, 1.03]	T
Total events	1509		1817				
Heterogeneity: Not applicab	ole						
Test for overall effect: Z = 0.	19 (P =	0.85)					
Total (95% CI)		3800		4757	100.0%	1.03 [1.00, 1.05]	•
Total events	2903		3486				a statistic concerning the state of the
Heterogeneity: Chi# = 49.88	df = 2	0 (P = 0)	0002) P	= 60%		-	- <del>1 1 1 1 1</del>
Test for overall effect: Z = 1	97 (P =	0.00			NG 201		05 07 1 15 2
Test for subgroup difference		U.U.D.I					0.0 0.1 1 1.0 2

Fig. 4 Subgroup analysis of recanalization success based on different sample size

stages following surgery [47]. An alternative perspective suggests that surgery under anesthesia offers enhanced intraoperative control and the capacity to address complications effectively, thereby facilitating superior longterm neurological recovery. Conversely, patients not under GA tended to experience swifter recovery in the immediate postoperative period, evident in their superior NIHSS scores within 24 h of surgery. This phenomenon could be partially attributed to the transient neurological depression induced by anesthetic drugs, although such effects might not endure over longer durations [48]. However, further research may be necessary to definitively ascertain the exact cause. In addition, for high-risk procedures such as IAT in patients with acute ischemic stroke, the volume of surgery performed by a medical center may have a certain impact on the patient's outcomes. A large number of studies have shown that there is a certain relationship between the volume of complex and high-risk surgeries and outcomes [49]. High-level medical centers may have more resources, more advanced equipment, and more experienced medical teams. Medical staff in low-sample centers may lack experience and training. This may have a certain impact on the results, so we also adopted subgroup analysis, meta-regression, and TSA to minimize this impact.

Propofol stands out as the predominant intravenous anesthetic in contemporary anesthesia practice. It mitigates post-ischemic neuronal damage through a variety of mechanisms, including the activation of GABAA receptors, exertion of antioxidant effects, reduction of brain mitochondrial membrane permeability, and augmentation of glutamate uptake. Additionally, Propofol diminishes cerebral blood flow (CBF), intracranial pressure, and cerebral oxygen metabolic rate, making it the preferred anesthetic for neurosurgical procedures [50]. Ketamine, an NMDA receptor antagonist, offers a distinct profile in that while it elevates CBF, its impact on overall cerebral oxygen metabolic rate remains minimal. Its neuroprotective properties stem from its ability to thwart the excitotoxic effects of glutamate, a mechanism supported by findings from in vitro and animal studies [51]. Moreover, intravenous lidocaine has emerged as another agent with demonstrated neuroprotective effects in the context of hypoxia-ischemia. This effect is likely attributed to its inhibition of sodium uptake and reduction of neuroinflammation, as evidenced by both in vitro and animal research [52]. In summary, the variance in prognosis observed among different anesthesia methods following IAT treatment for ischemic stroke may, in part, be attributed to the neuroprotective effects of anesthetic drugs. GA could potentially confer neuroprotective benefits in certain disease states linked to cerebral ischemia. While animal experiments offer robust support for this notion, clinical evidence remains scarce. Hence, future research should prioritize investigating the neuroprotective properties of various anesthetic drugs. Subsequently, these findings can inform the selection of safer medications tailored to specific clinical patients or scenarios. This approach aims to fulfill the demands for swift recovery and personalized diagnosis and treatment.

Although our study included a large number of existing studies, it still has the following limitations: First, half of the included literature were non-randomized controlled studies, which may put us at a disadvantage in terms of the level of evidence. Second, we were unable to conduct subgroup analyzes according to different stroke conditions and different anesthetic drugs due to limitations of baseline data and few studies reporting specific anesthetic regimens. Finally, there are differences in the actual implementation of specific anesthesia methods among multiple centers, and this irremovable bias may also have an impact on outcomes. In particular, different anesthesiologists and neurologists may have personal preferences, which cannot be eliminated.

# Conclusion

Utilizing IAT without GA presents clear benefits for patients suffering from ischemic stroke. These advantages include a reduced risk of mortality, an increased rate of successful reperfusion, and shorter hospital stays. Regarding neurological outcomes, patients undergoing IAT without GA tend to experience fewer short-term postoperative deficits. However, when considering longterm neurological outcomes, GA may yield superior results. In addition, GA may have a smaller risk of vascular perforation and a higher risk of intracranial hemorrhage after surgery. Therefore, for the anesthesia plan of IAT in the future, excluding unstable or critically dangerous patients, patients with high risk of aspiration pneumonia and other routine non-special situations, we can consider more non-intubation GA methods, such as MAC and CS. When conditions permit, the anesthesia plan should be fully evaluated and discussed by anesthesiologists and neurologists, and the decision should be made after the three parties have discussed and educated the patient's family.

#### **Supplementary Information**

The online version contains supplementary material available at https://doi.org/10.1186/s12871-024-02633-3.

Supplementary Material 1
Supplementary Material 2
Supplementary Material 3
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#### Acknowledgements

Not applicable.

#### Author contributions

Huijun Chen and Ximin He designed the study. Zekun Lang was responsible for the main statistical analysis. Huijun Chen and Xing Yang were responsible for most of the manuscript writing, Lei Zhang and Ximin He were respectively responsible for part of the manuscript writing, data extraction and literature screening. Mao Liao is responsible for the organization and production of some charts. Ximin He reviewed the full text and made relevant suggestions.

#### Funding

No funding.

#### Data availability

All data generated or analysed during this study are included in this published article [and its supplementary information files].

#### Declarations

**Ethical approval and consent to participate** Not applicable.

#### **Consent to publish**

Not applicable.

#### **Competing interests**

The authors declare no competing interests.

Received: 30 April 2024 / Accepted: 11 July 2024 Published online: 18 July 2024

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