

Video vs. direct laryngoscopy for adult surgical and intensive care unit patients requiring tracheal intubation: a systematic review and meta-analysis of randomized controlled trials

M. VARGAS¹, G. SERVILLO¹, P. BUONANNO¹, C. IACOVAZZO¹, A. MARRA¹, G. PUTENSEN-HIMMER², S. EHRENTAUT², L. BALL³, N. PATRONITI⁴, P. PELOSI⁴, C. PUTENSEN²

¹Department of Neurosciences, Reproductive and Odontostomatological Sciences, University of Naples "Federico II", Naples, Italy

²Department of Anesthesiology and Intensive Care Medicine, University Hospital Bonn, Bonn, Germany

³San Martino Policlinico Hospital, IRCCS for Oncology and Neurosciences, Genoa, Italy

⁴Department of Surgical Sciences and Integrated Diagnostics, San Martino Policlinico Hospital, IRCCS for Oncology, University of Genoa, Genoa, Italy

Abstract. – **OBJECTIVE:** This systematic review and meta-analysis aimed to determine whether a specific videolaryngoscopy technique is superior to standard direct laryngoscopy using a Macintosh blade to reduce the risk of difficult intubation in surgical and intensive care unit patients.

MATERIALS AND METHODS: We identified all randomized controlled trials comparing videolaryngoscopes (VLSs) to direct laryngoscopy in the Cochrane Central Register of Controlled Trials, MEDLINE, and EMBASE (from inception to April 2020). The primary outcome was difficult intubation in adult surgical and intensive care unit patients. Secondary outcomes were successful intubation at the first attempt, airway trauma, sore throat, hoarseness, hypoxia, and mortality.

RESULTS: We included 97 randomized controlled trials to evaluate 12775 patients. A high risk of bias was found in at least 50% of the included studies for each outcome. VLSs reduced the risk of difficult intubation compared to direct Macintosh laryngoscopy (RR 0.48, 95% CI from 0.35 to 0.65). VLSs increased the rate of successful intubation at the first attempt when compared to direct Macintosh laryngoscopy (RR 1.03, 95% CI from 1.00 to 1.07). Lower risks of airway trauma were found with VLSs (RR 0.69, 95% CI from 0.55 to 0.86). A decreased risk of hoarseness was associated with the use of VLSs (RR 0.67, 95% CI from 0.54 to 0.83). In addition, VLSs did not significantly reduce the risk of hypoxia compared with direct laryngoscopy (RR 0.83, 95% CI from 0.60 to 1.16).

CONCLUSIONS: In this systematic review and meta-analysis, we found that the use of VLSs reduced the risk of difficult intubation and slightly increased the ratio of successful intubation at the first attempt among adult patients.

Key Words:

Videolaryngoscope, Direct laryngoscopy, Randomized controlled trial, Operation room.

Abbreviations

VLS: videolaryngoscope; RCT: randomized controlled trials; ICU: intensive care unit; OR: operation room; RR: risk ratio.

Introduction

Tracheal intubation is used for airway management in more than 1.1 million patients per year¹. Intubation with direct laryngoscopy is a complex procedure requiring flexing the lower cervical spine and extending the upper cervical spine to create a "line of sight" and a Macintosh blade to retract the tongue to enable passage of a tracheal tube¹. Difficult intubation occurs in 1% to 6% of endotracheal intubation while intubation failure occurs in 0.1% to 0.3%, of them and both are associated with complications such as desaturation, hyper- and hypotension, airway damage, and death².

Many airway characteristics have been associated with difficult intubation, such as limited mouth opening, limited mandibular protrusion, narrow dental arch, decreased thyromental distance, Mallampati class 3 or 4, decreased submandibular compliance, decreased sterno-mental distance, limited head and upper neck extension, and increased neck circumference^{1,2}. However, a large observational study showed that 93% of difficult intubations are unpredictable, even using predictive tests¹.

In recent years, videolaryngoscopes (VLSs) have been increasingly used to manage difficult airways³. VLSs rely on fiberoptic or digital technology to transmit an image from the tip of the laryngoscope to an eyepiece or videomonitor, thereby giving the intubator a better indirect view of the glottis independent of the line of sight¹. Different types of VLSs have been extensively studied in the operation room (OR) and the intensive care unit (ICU) with the assumption of improving the success of tracheal intubation². VLSs may help manage patients with a predicted or known difficult airway since their use is likely to improve the glottic view and reduce the number of laryngoscopies in which the glottis cannot be visualized². Previous systematic reviews comparing VLSs with direct laryngoscopy suggested that VLSs may reduce the number of difficult intubations in patients presenting with a difficult airway^{1,2}. However, there are conflicting data in the literature regarding whether the use of VLSs results in increased success of first-pass intubation attempts, decreased complications, or improved clinical outcomes³. These conflicting reports could depend on the high level of performance bias due to the lack of blinding and the high heterogeneity of the studies. Therefore, it is unclear whether VLSs should be the new standard of care for all tracheal intubations². Our review aims to update the Cochrane review by Lewis et al² with the more recent randomized controlled trials (RCTs) evaluating the use of VLSs vs. Macintosh published in the last few years². The primary outcome was difficult intubation. Difficult intubation was defined as intubation 1) not achieved with \geq two attempts, 2) lasting longer than 60 s, or 3) tried with an alternative device. Furthermore, as secondary outcomes, we evaluated whether VLSs improve successful intubation at the first attempts and reduce the risk of airway trauma, sore throat, hoarseness, hypoxia, and mortality.

Materials and Methods

The protocol for this meta-analysis was registered at PROSPERO (CRD42019107564) after the commencement of the systematic review process.

Data Sources and Searches

We searched for all RCTs comparing VLSs to direct laryngoscopy regarding difficult intubation, successful first-attempt intubation, airway trauma, sore throat, hoarseness, and hypoxia in adult patients. The electronic search strategy applied standard filters for the identification of RCTs. Databases searched were the Cochrane Central Register of Controlled Trials, MEDLINE, and EMBASE (from inception to April 2020). However, we did not apply any language restrictions. Our search included the following keywords: intubation, videolaryngoscopy, videolaryngoscope, anesthesia, critical care, intensive care, critically ill, emergency care, airway trauma, and randomized controlled trial. In addition to the electronic search, we checked out cross-references from original articles and reviews. No ethical approval will be needed because data from previously published studies in which informed consent was obtained by primary investigators will be retrieved and analyzed.

Selection of Studies

We restricted the analysis to RCTs and excluded study designs containing co-interventions unequally applied to the treatment and control group, non-randomized, or cross-over trials. Additionally, we considered for inclusion all RCTs reporting difficult intubation, successful first-attempt intubation, airway trauma, sore throat, hoarseness, and hypoxia as predefined endpoints and comparing VLS to direct laryngoscopy using a Macintosh blade in adult patients. Furthermore, we excluded RCTs performed in emergency out-of-hospital settings, in pediatric patients, and using direct laryngoscopes other than Macintosh.

Outcome Measures

The primary outcome was difficult intubation. Difficult intubation was defined as intubation 1) not achieved with \geq two attempts, 2) lasting longer than 60 s, or 3) tried with an alternative device. Secondary outcomes included successful intubation at the first attempt, airway trauma, sore throat, hoarseness, hypoxia, and mortality. [Supplementary Materials 1](#) show the outcome definitions.

Subgroup Analysis

We performed subgroup analyses to assess the impact on primary and secondary outcomes:

1. Type of VLS devices: Airtraq (Prodol Meditec, Guecho, Spain), C and V-Mac (Karl Storz Endoscopy, Tuttlingen, Germany), GlideScope (Verathon Inc., Bothell, Washington, U.S.A.), McGrath (Aircraft Medical Ltd., Edinburgh, UK), Pentax AWS (Pentax Corporation, Tokyo, Japan), Truview (Truphatek International Ltd., Netanya, Israel), X-lite (Teleflex Medical Europe Ltd, Westmeath, Ireland). Table II in [Supplementary Materials 2](#) show the main characteristics of the different VLS devices.
2. Clinical scenario: operation room or intensive care unit
3. Operator experience: experienced or less experienced intubator
4. Predicted, not predicted, and simulated difficult airway
5. Based on the previous studies, we defined “experienced intubators” as those who had performed more than 20 successful intubations with each device².

Data Extraction and Quality Assessment

Two pairs of independent reviewers performed an initial selection through the screening of titles and abstracts (MV and PB, PP and CP). Each reviewer screened citations to identify further RCTs to include in our analysis. For detailed evaluation, a full-text copy of all possibly relevant studies was obtained. Each study’s data were extracted independently by paired reviewers (MV and PB, PP and CP) using a pre-standardized data extraction form. One pair of reviewers (GS and LB) was not informed about the authors, journal, institutional affiliations, and date of publication. Another reviewer checked the data extracted from the publications for accuracy. Furthermore, we used the Cochrane risk of bias tool to assess the quality of study design and the extent of potential bias^{5,6} by considering the following domains: random sequence generation, allocation concealment, blinding of participants, personnel and outcomes assessors, incomplete outcome data, selective outcomes reporting, baseline characteristics, and funding resources. Two reviewers (MV and PB) independently used these criteria to abstract trial quality. We resolved any disagreements by consensus consultation with a third reviewer (LB) if needed.

Qualitative Analysis

The risk of bias was assessed using the Cochrane Risk of Bias Tool for RCTs and the Risk of Bias Instrument for Non-Randomized Studies of Interventions (ROBINS-2)⁵. The certainty of the evidence was assessed using the GRADE approach⁶. According to GRADE, the body of evidence for each outcome was assessed for factors that may reduce or increase it⁶. GRADE summary of findings and tables were developed with GRADEpro GDT software (McMaster University, 2015. Developed by Evidence Prime, Inc. Available at: <https://gradepr.org/>).

Quantitative Analysis

The meta-analysis was conducted according to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines ([Supplementary Materials 3](#)). Meta-analysis was performed with mixed random effects using DerSimonian and Laird methods. Results were graphically represented using Forest plot graphs. According to an intention-to-treat principle, the relative risk (RR) and 95% CI for each outcome were separately calculated for each trial, pooling data when needed. The choice to use RRs was driven by the design of a meta-analysis based on RCTs. Tau² defined the between-studies variance. The difference in treatment effect estimates between the treatment groups for each hypothesis was tested using a two-sided Z test with statistical significance considered at a *p*-value of less than 0.05. The homogeneity assumption was checked with a Q test with a degree of freedom (df) equal to the number of analyzed studies minus 1. The heterogeneity was measured by the I² metric, which describes the percentage of total variation across studies that is due to heterogeneity rather than chance. I² was calculated as $I^2 = 100\% \cdot (Q - df)/Q$, where Q is Cochran’s heterogeneity statistic, and df is the degrees of freedom. A value of 0% indicates no observed heterogeneity, and larger values show increasing heterogeneity. Analyses were conducted with OpenMetaAnalyst (version 6) and SPSS version 20 (IBM SPSS Corp., Armonk, NY, USA)). To evaluate potential publication bias, a weighted linear regression was used, with the natural log of the RR as the dependent variable and the inverse of the total sample size as the independent variable. This modified Macaskill’s test gives more balanced type I error rates in the tail probability areas than other publication bias tests. The number needed to treat (NNT) was calculated as the inverse of the absolute risk re-

duction (ARR) expressed as a decimal. Cumulative meta-analyses of RCTs are at risk of yielding random errors due to sparse data and repetitive testing of accumulating data⁷. Trial sequential analysis (TSA) depends on the quantification of the required information size (RIS) (i.e., optimal information size). TSA was undertaken using TSA 0.9 beta software if the number of included trials was more than five. The RIS was estimated using relative risk reduction and heterogeneity adjusted information size for dichotomous outcomes. The result was confirmed as a true positive if the cumulative Z-curve surpassed the Lan-DeMets trial sequential monitoring boundary or reached the RIS above the conventional significance level line ($Z = 1.96$). Moreover, the result was confirmed as a true negative if the cumulative Z-curve reached the futility boundary or reached the RIS below the conventional significance level line ($Z = -1.96$). TSA adjusted 95% CIs were also presented.

Data Synthesis and Analysis

We classified the following comparisons: pooled VLS devices vs. direct laryngoscopy (Macintosh laryngoscope), Airtraq vs. direct laryngoscopy, C and V-Mac vs. direct laryngoscopy, GlideScope vs. direct laryngoscopy, McGrath vs. direct laryngoscopy, Pentax AWS vs. direct laryngoscopy, Truview vs. direct laryngoscopy, and X-lite vs. direct laryngoscopy.

Results

In this systematic review and meta-analysis, we included 97 RCTs, evaluating 12775 patients (Figure 1)⁸⁻¹⁰⁴. VLSs were investigated in elective surgical patients in 92 studies^{8,10,12-35,37-48,49-51,53-78,81-91,93-99,101-104}, in the ICU in five studies^{36,52,79,92,100} and in the Emergency Department in four RCTs^{11,49,80,92}. We found eight RCTs comparing Airtraq vs. direct laryngoscopy^{9-11,13,14,16,25,26,29,31,32,36,41,44,48,53,56,60,62,66,69,70-73,75,76,78,81,85,86,89,91,92,95,96}, 19 RCTs comparing McGrath vs. direct laryngoscopy^{17,20,34,35,42,43,47,51,52,54,58,68,74,77,84,88,93,94,96}, 18 RCTs comparing Pentax AWS vs. direct laryngoscopy^{8,15,18,33,39,40,49,50,55,60,62,63,67,82,83,85,90}, two RCTs comparing Truview vs. direct laryngoscopy^{30,61}, and two RCTs comparing X-lite vs. direct laryngoscopy^{21,22}. We found 11 RCTs comparing more than one VLS with direct laryngoscopy^{20,30,53,54,60,62,65,75,76,85,96}.

The Cochrane risks of bias tool were used to assess the quality of the study design and the extent

of potential bias (Figure 2 and [Supplementary Materials 4](#)). We found that 42 out of 60 studies for the primary outcome, 54 out of 80 studies for successful intubation at the first attempt, 36 out of 51 studies for airway trauma, 20 out of 26 studies for sore throat, 6 out of 13 studies for hoarseness, 8 out of 17 studies for sore throat, and 2 out of 4 studies for mortality had a high risk of bias.

The risk of difficult intubation was significantly lower when using VLSs compared to direct Macintosh laryngoscopy (RR 0.48, 95% CI from 0.35 to 0.65, $I^2 = 16\%$) (NNT = 25, 95% CI = 19.8-32.3) (Figure 3).

The risk of difficult intubation was reduced by C and V-MAC (RR 0.40, 95% CI from 0.20 to 0.79, $I^2 = 0\%$) and Airtraq (RR 0.45, 95% CI from 0.23 to 0.87, $I^2 = 67\%$) but not by the other VLSs ([Supplementary Materials 5](#)) (GlideScope vs. direct Macintosh laryngoscopy: 23 studies, RR 0.61, 95% CI from 0.36 to 1.05, $I^2 = 0\%$) (McGrath vs. direct Macintosh laryngoscopy: 16 studies, RR 0.68, 95% CI from 0.27 to 1.69, $I^2 = 53\%$) (Pentax AWS vs. direct Macintosh laryngoscopy: 10 studies, RR 0.35, 95% CI from 0.12 to 1.05, $I^2 = 29\%$) (Truview vs. direct Macintosh laryngoscopy: 3 studies, RR 0.99, 95% CI from 0.28 to 3.49, $I^2 = 0\%$) (X-lite vs. direct Macintosh laryngoscopy: 2 studies, RR 0.25, 95% CI from 0.03 to 2.48, $I^2 = 0\%$).

VLSs reduced the risk of difficult intubation in the OR (RR 0.46, 95% CI from 0.33 to 0.64, $I^2 = 16.5\%$), in predicted and simulated difficult airways (RR 0.32, 95% CI from 0.17 to 0.59, $I^2 = 3.4\%$ and RR 0.23, 95% CI from 0.11 to 0.50, $I^2 = 0\%$, respectively) ([Supplementary Materials 5](#)). VLSs, when used by an experienced operator, reduced the risk of difficult intubations (RR 0.48, 95% CI from 0.30 to 0.77, $I^2 = 0\%$) ([Supplementary Materials 6](#)). Excluding the simulated of funded RCTs, the result for the primary outcome persisted ([Supplementary Materials 6](#)). By including 5521 patents, the Z-curve crosses the TSA monitoring boundary, and the RIS (=1627) for this specific TSA-analysis is conclusive in favor of VLSs (Figure 4).

VLSs increased the rate of successful intubation at the first attempt when compared to direct Macintosh laryngoscopy (RR 1.03, 95% CI from 1.00 to 1.07, $I^2 = 79\%$) (NNT = 18, 95%CI = 13.9-26) (Figure 5). The TSA adjusted 95% CI ranged from 0.77 to 0.81. The cumulative Z-curve crossed neither the conventional boundary for benefit nor the trial sequential futility boundary for benefit, suggesting that the current evidence

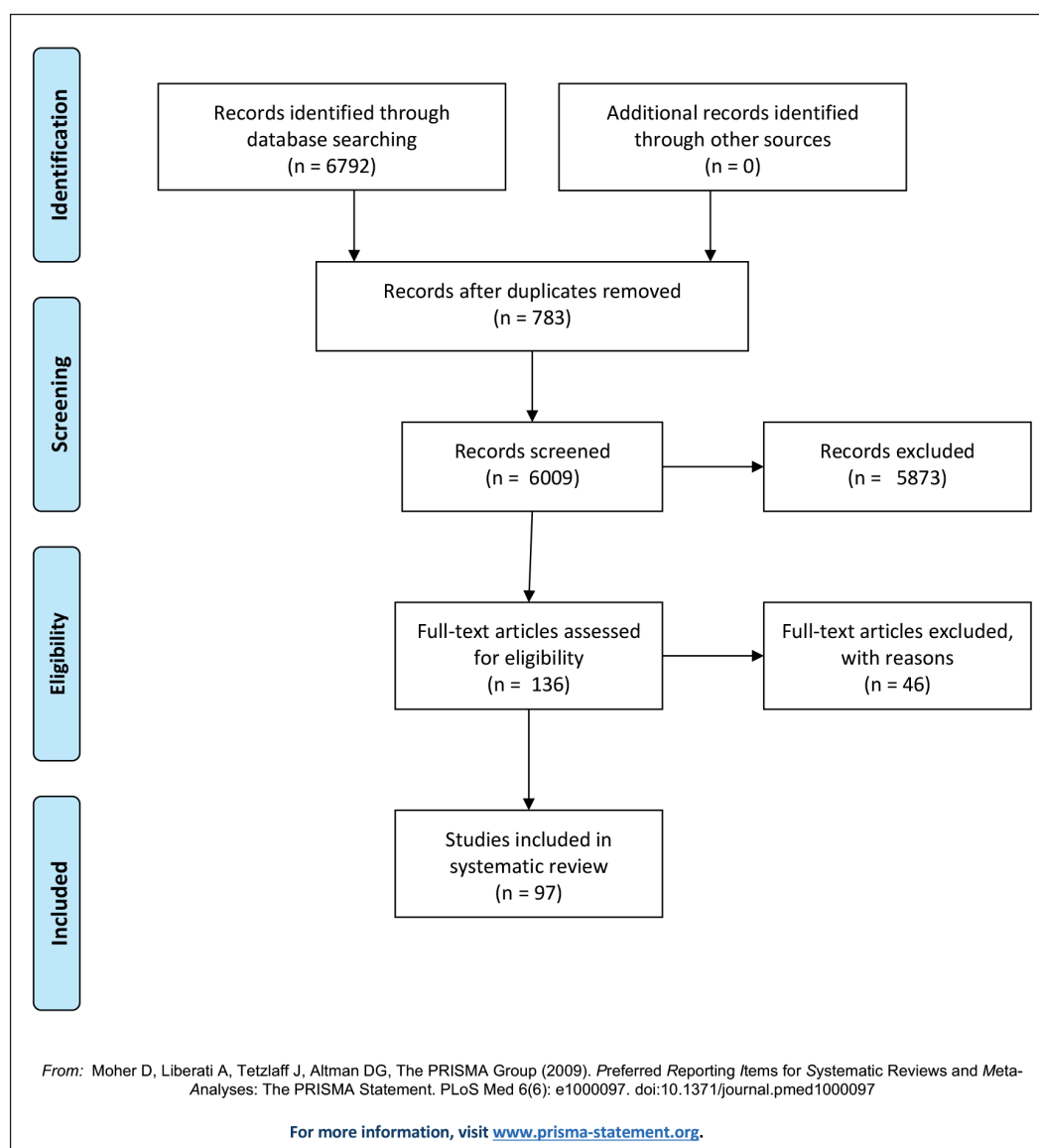


Figure 1. PRISMA flow chart of included studies.

was inconclusive (**Supplementary Materials 7**). Different analyses performed according to VLS devices, clinical scenario, operator experience, and difficult airways did not show significant results (**Supplementary Materials 8**).

Lower risks of airway trauma were found with VLSs (RR 0.69, 95% CI from 0.55 to 0.86, $I^2 = 7.6\%$) (NNT = 41, 95%CI = 26.7-87.6) (**Supplementary Materials 9**). This result was mainly due to McGrath VLSs, which significantly decreased the risk of airway trauma compared to direct laryngoscopy (RR 0.54, 95% CI from 0.36 to 0.81, $I^2 = 0\%$) (**Supplementary Materials 10**). Airway trauma was reduced by using VLSs in OR (RR 0.69,

95% CI from 0.54 to 0.87, $I^2 = 0\%$), in predicted not difficult airway (RR 0.57, 95% CI from 0.38 to 0.85, $I^2 = 10\%$), and in simulated difficult airway (RR 0.60, 95% CI from 0.39 to 0.90, $I^2 = 18\%$) (**Supplementary Materials 10**). The TSA adjusted 95% CI ranged from 0.15 to 0.42. The TSA result showed that 3457 of the RIS of 5882 patients was accrued. The cumulative Z-curve crossed the conventional boundary for the benefit and crossed the trial sequential monitoring boundary for benefit (**Supplementary Materials 11**), indicating firm evidence of VLSs for airway trauma.

Analysis for sore throat did not reveal the advantage of using VLS compared to direct Macin-

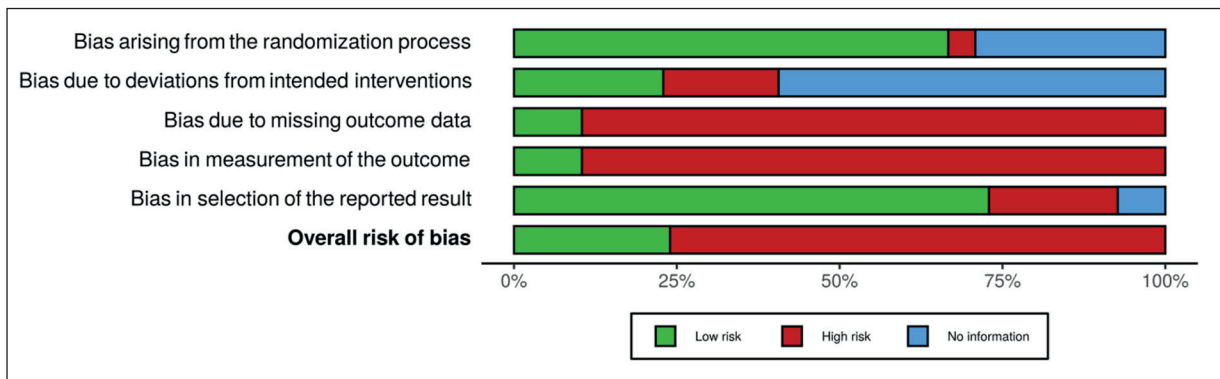


Figure 2. Risk of bias summary. Green represents a low risk of bias, yellow some concerns, and red a high risk of bias.

tosh laryngoscopy (RR 0.85, 95% CI from 0.72 to 1.01, $I^2 = 50\%$) (NNT = 14, 95%CI = 9.4-24.4) (Supplementary Materials 12). The sub-group analyses did not show any significant results (Supplementary Materials 13). Different analyses performed according to the type of VLS devices, operator experience, and difficult airways did not show significant results (Supplementary Materials 12). The TSA adjusted 95% CI ranged from 0.43 to 0.64. The cumulative Z-curve crossed the conventional boundary for the benefit and crossed the trial sequential monitoring boundary for benefit (Supplementary Materials 14 and 15), indicating that this evidence was conclusive.

A decreased risk of hoarseness was associated with the use of VLS (RR 0.675, 95% CI from 0.54 to 0.84, $I^2 = 20\%$) (NNT = 10, 95% CI = 6.5-15.6) (Supplementary Materials 16). Risk of hoarseness was only reduced with Glidescope VLS (RR 0.52, 95% CI from 0.39 to 0.69, $I^2 = 0\%$) (Supplementary Materials 17). The TSA adjusted 95% CI ranged from 0.00 to 0.43. The cumulative Z-curve crossed neither the conventional boundary for benefit nor the trial sequential futility boundary for benefit, suggesting that the current evidence was inconclusive (Supplementary Materials 14 and 18).

VLS was not associated with a reduced risk of hypoxia compared to direct Macintosh laryngoscopy (RR 0.83, 95% CI from 0.60 to 1.16, $I^2 = 0\%$) (NNT = 29, 95%CI = 18.7-63.2) (Supplementary Materials 19 and 20). The TSA adjusted 95% CI ranged from 0.34 to 0.73. The alpha boundary for hypoxia was not performed because of limited extant information (4%) (Supplementary Materials 14 and 21).

The risk of mortality was not reduced by using VLS compared to direct Macintosh laryngoscopy (RR 1.03, 95% CI from 0.69 to 1.516, $I^2 = 0\%$)

(Supplementary Materials 22 and 23). The TSA adjusted 95% CI ranged from 0.00 to 0.60. The cumulative Z-curve crossed neither the conventional boundary for benefit nor the trial sequential futility boundary for benefit, suggesting that the current evidence was inconclusive (Supplementary Materials 14 and 24).

Overall, evidence was qualified using GRADE for RCTs (Figure 6). High-quality evidence was found for difficult intubation, successful intubation at the first attempt, airway trauma, and sore throat. RCTs' level of evidence was downgraded due to the high risk of bias mainly caused by allocation concealment, blinding of participants and personnel, and funding in outcome assessment in most studies.

Discussion

This systematic review and meta-analysis in adult patients show a reduced risk of difficult intubation with VLSs than direct Macintosh laryngoscopy. The risk of difficult intubation was reduced using C-and V-MAC and Airtraq devices when used by experienced physicians in predicted or simulated difficult airways in the OR and in the ICU setting. As qualified by GRADE, the quality of evidence for difficult intubation was high, suggesting that further research is unlikely to impact our estimated effect or change it significantly. VLS slightly increased the successful intubation at the first attempt and reduced the risk of airway trauma and hoarseness. At the same time, it did not affect the risk of sore throat, hypoxia, or mortality. Even the quality of evidence qualified by GRADE for successful intubation at the first attempt, airway trauma, and sore throat was high, suggesting that further research is very unlikely to have a significant impact on our estimated effect.

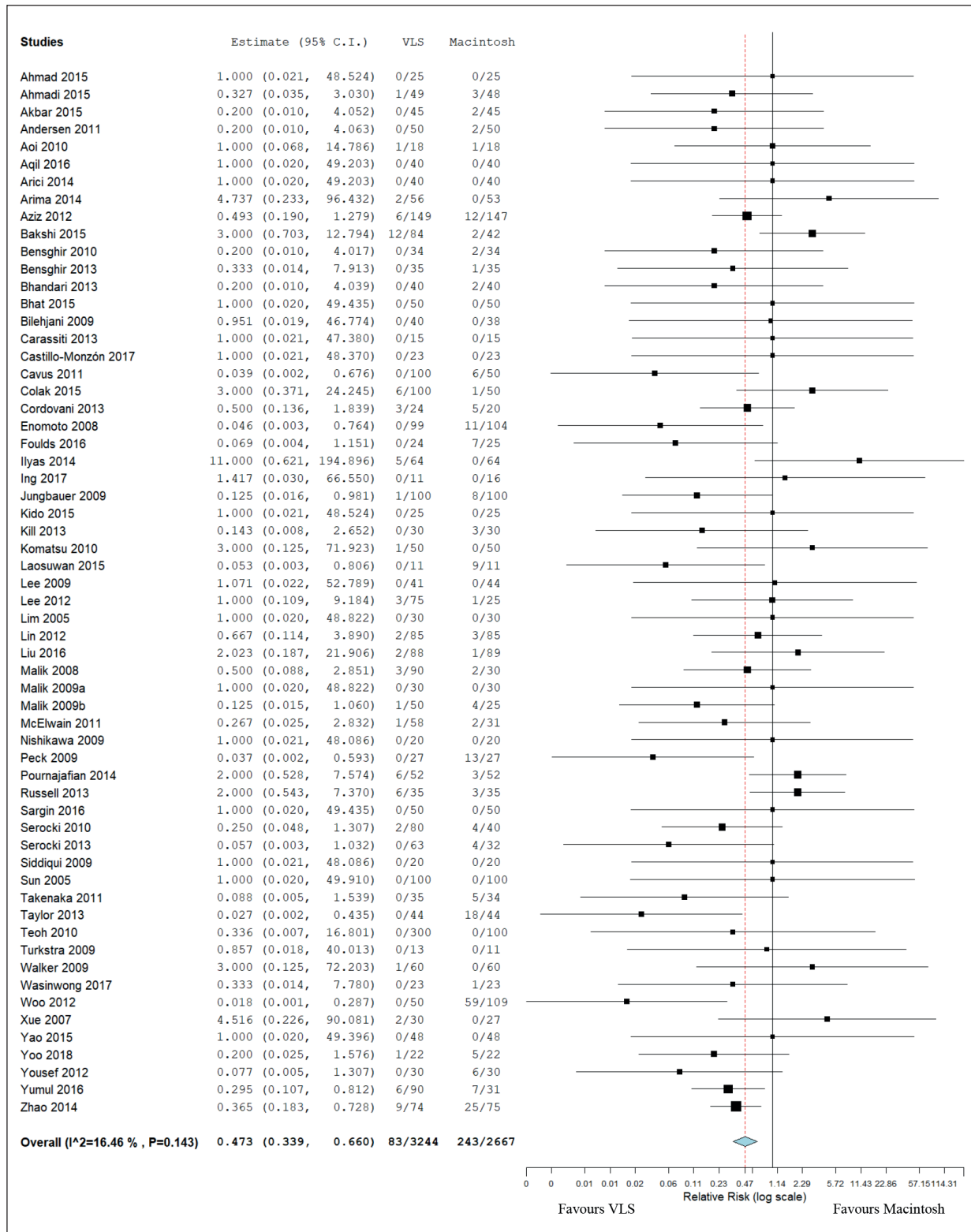


Figure 3. Forest plot for difficult intubation of VLSs vs. Macintosh. VLS: videolaryngoscope. C.I.: confidence interval.

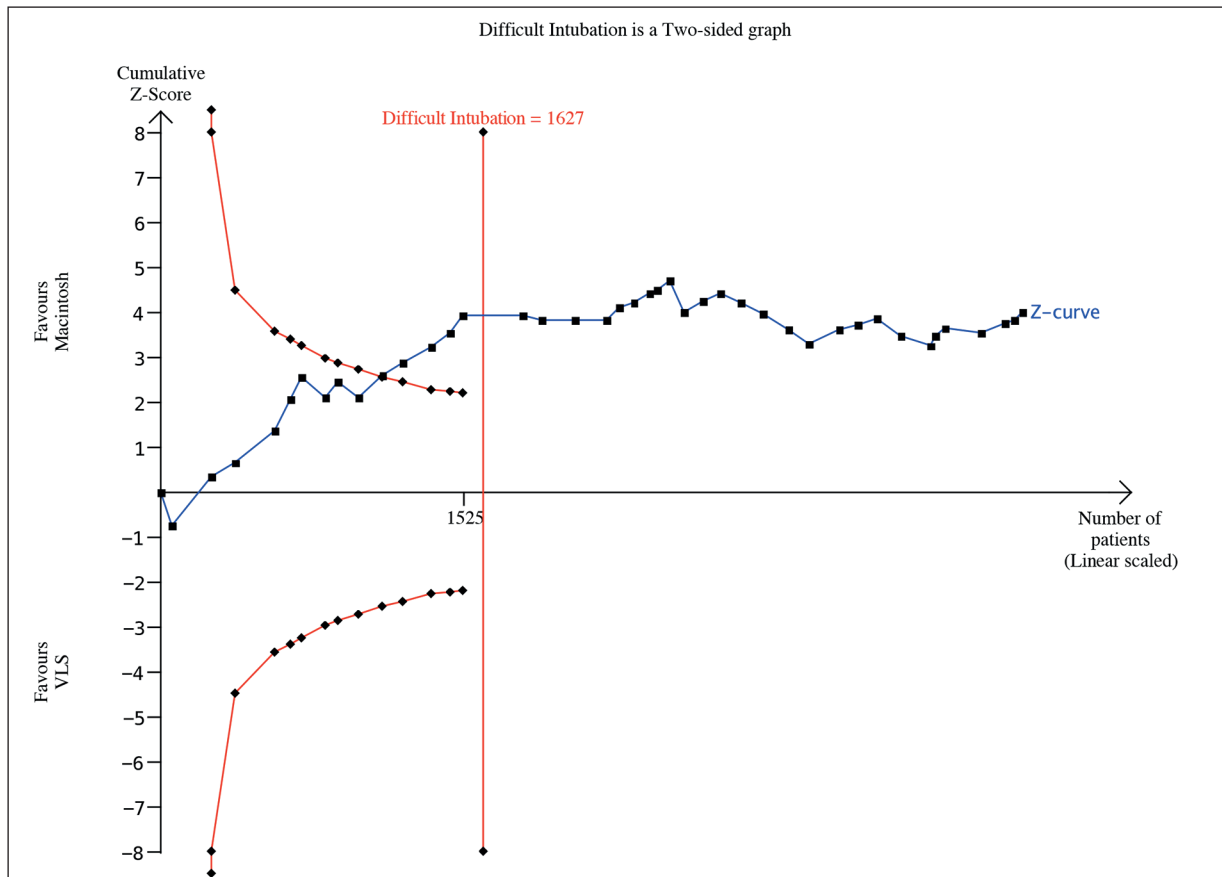


Figure 4. Trial sequential analysis for difficult intubation of VLSs vs. Macintosh. TSA was performed with Relative Risk and Random-effects (Der-Simonian and Laird). Zero-event Trials are not included. Pooled Effect: 0.41 (C.I.: 0.26 to 0.63) based on conventional 95%. Inconsistency (I^2): 0.46; Diversity (D^2): 0.57. Boundaries: Type 1 Error: 5.0%, Alpha Spending: O'Brien-Fleming, Power: 80.0%.

Previous systematic reviews and meta-analysis investigating the use of VLSs vs. direct Macintosh laryngoscopy included cross-over studies^{2,105}; were restricted to patients with predicted difficult airways¹⁰⁶, cervical immobilization¹⁰⁷, or receiving treatment in the ICU or emergency department¹⁰⁸⁻¹¹³; were limited to procedures performed by trained anesthetists¹¹⁴; were limited to pediatric¹¹⁵ and neonatal patients¹¹⁶; excluded particular VLSs types such as the Airtraq and Truview device^{2,106}. Conversely, our meta-analysis included the most recent high-quality RCTs in adult patients; moreover, we performed subgroup analyses to investigate the different VLS types, including Airtraq and Truview devices.

Among the different VLS types, only the Airtraq, C-and V-Mac devices reduced the risk of difficult intubation. Compared to Macintosh laryngoscopy, the reduced risk of difficult intubation with the Airtraq has been reported in patients with cervical spine immobilization¹¹⁴. This benefit

may be explained by the more pronounced curvature and the integral tube guidance channel of the Airtraq blade, which does not require alignment of oral, pharyngeal, and tracheal axes^{115,116} and better positioning of optical components^{100, 115,116}. The Pentax AWS device has a similar curvature and guides channel, but the tube may strike the epiglottis and may be challenging to advance, making intubation more difficult¹¹⁷. Reduced risk of difficult intubation observed with the C-and V-Mac devices using a classic Macintosh blade was unexpected. Physicians experienced with direct tracheal intubation, trying subconsciously to bring the axes in one line when intubating, may have fewer difficulties and do not need new skill requirements when using VLS devices with a classic shaped Macintosh blade like the C-and V-Mac¹⁰⁷ as compared to GlideScope, McGrath, and Truview devices¹⁰⁷. VLSs reduced the risk of difficult intubations in patients with predicted or simulated difficult airways, in line with previous reports^{2,105-107}.

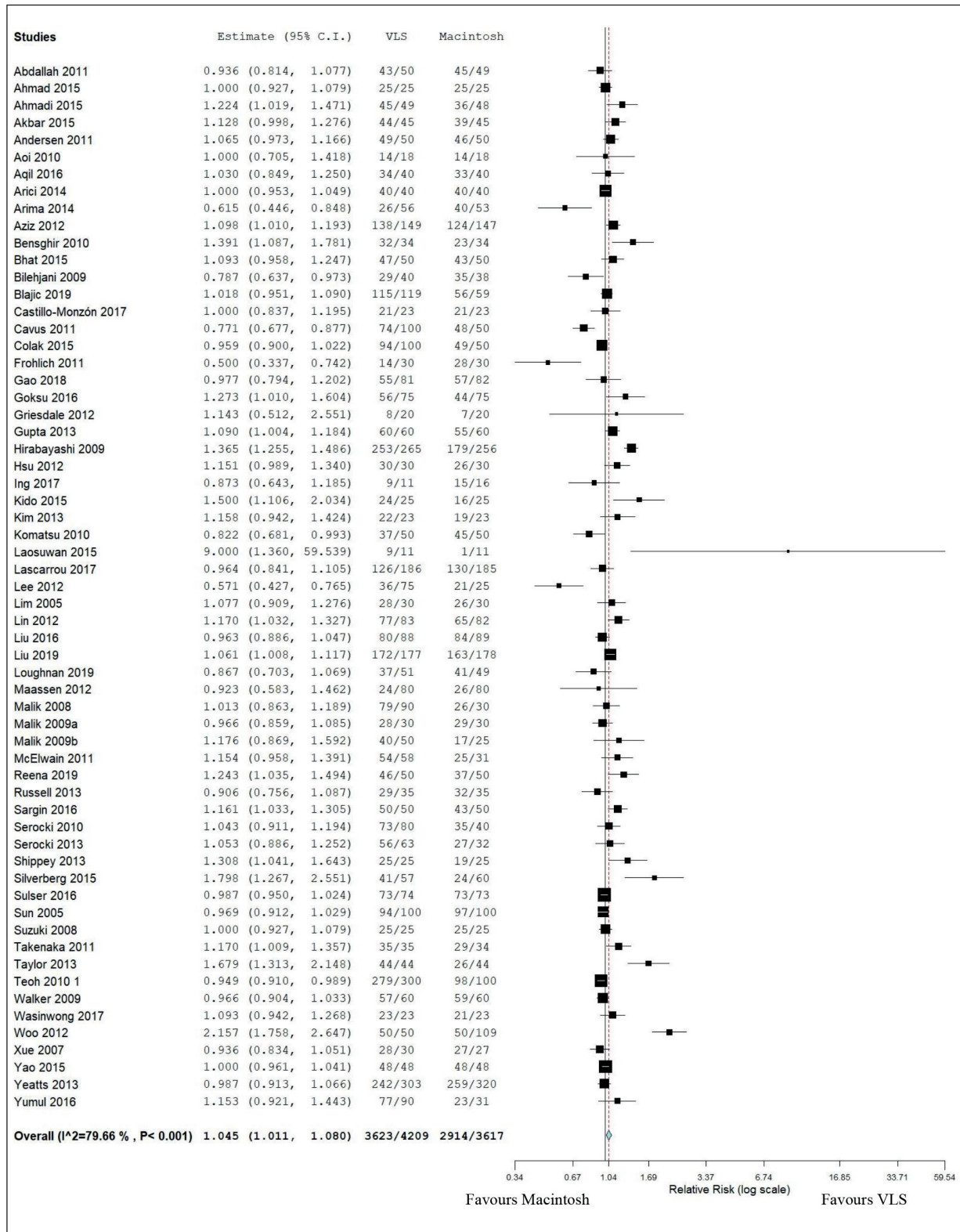


Figure 5. Forest plot for successful at first attempts of VLSs vs. Macintosh. VLS: videolaryngoscope. C.I.: confidence interval.

In contrast with Lewis et al², this meta-analysis found that VLSs increased the rate of suc-

cessful intubation at the first attempt compared to direct Macintosh laryngoscopy. Several patients

A meta-analysis on videolaryngoscope vs. direct laryngoscopy

Author(s): Question: Videolaryngoscopy compared to Direct laryngoscopy for tracheal intubation Setting: operation room Bibliography:												
No. of studies	Study design	Certainty assessment					No. of patients		Effect		Certainty	Importance
		Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Videolaryngoscopy	Direct (laryngoscopy)	Relative (95% CI)	Absolute (95% CI)		
Difficult intubation												
60	randomised trials	serious ^a	not serious	not serious	not serious	strong association	83/3244 (2.6%)	243/2667 (9.1%)	RR 0.47 (0.34 to 0.66)	48 fewer per 1,000 (from 60 fewer to 31 fewer)	⊕⊕⊕⊕ HIGH	CRITICAL
Successful intubation at the first attempt												
80	randomised trials	serious ^b	not serious	not serious	not serious	strong association	3623/4209 (86.1%)	2914/3617 (80.6%)	RR 1.04 (1.01 to 1.08)	32 more per 1,000 (from 8 more to 64 more)	⊕⊕⊕⊕ HIGH	CRITICAL
Airway trauma												
51	randomised trials	serious ^c	not serious	not serious	not serious	strong association	181/3226 (5.6%)	214/2656 (8.1%)	RR 0.69 (0.56 to 0.86)	25 fewer per 1,000 (from 35 fewer to 11 fewer)	⊕⊕⊕⊕ HIGH	CRITICAL
Sore throat												
26	randomised trials	serious ^d	not serious	not serious	not serious	strong association	371/1527 (24.3%)	407/1268 (32.1%)	RR 0.87 (0.73 to 1.04)	42 fewer per 1,000 (from 87 fewer to 13 more)	⊕⊕⊕⊕ HIGH	CRITICAL
							0.0%	0 fewer per 1,000 (from 0 fewer to 0 fewer)				
Hoarseness												
13	randomised trials	serious ^e	not serious	not serious	not serious	none	105/668 (15.7%)	163/614 (26.5%)	RR 0.67 (0.52 to 0.88)	88 fewer per 1,000 (from 127 fewer to 32 fewer)	⊕⊕⊕○ MODERATE	IMPORTANT

Figure 6. GRADE evidence profile for considered outcome.

and physician-related factors have been identified to be associated with first-attempt intubation failures, such as the anatomy of the supra- and infra-glottic airways, visibility of the glottis during laryngoscopy, device selection, pharmacologic choices, depth of anesthesia, adequate neuromuscular blockade, and experience^{2,119,122}. In addition, increasing time to intubate may be associated with increased intubation-related complications, such as hypoxemia. Using VLS, superior views of the glottis are often obtained but may not consistently translate into easier tracheal intubation and a higher rate of first intubation success¹⁹.

Our data suggest that VLS should be immediately available and that all physicians in OR and ICU, especially non-experienced physicians, should be trained in its use^{118,119}. With the high quality of evidence, our data demonstrated that the use of VLSs reduces the risk of airway trauma, probably because the use of VLSs may support the glottic view, allowing the application of less force during laryngoscopy¹²⁰⁻¹²².

This meta-analysis has several strengths. TSA was conclusive for the primary outcome in favor of VLS. According to the GRADE approach, we

found a high quality of evidence for 4 out of 7 outcomes.

Our findings should be interpreted regarding several limitations. First, large-scale RCTs, comparing VLSs with direct Macintosh laryngoscopy, are missing, and all RCTs included in this meta-analysis enrolled only a small number of patients. Second, the generalizability of our results may be limited since particular clinical scenarios, such as emergency or out-of-hospital settings, have not been sufficiently explored in the included RCT. Third, in the included RCTs, different criteria were used to define the physician's intubation expertise. Similar to previous meta-analyses^{2,105}, we considered a physician experienced after 20 successful intubations using both VLSs and direct laryngoscopy, while a systematic review, analyzing learning curves, suggested 50 successful intubations¹²³⁻¹²⁵. We were able to distinguish performance differences between VLS types. Fourth, although only high-quality RCTs, the first-attempt intubation success rate showed substantial heterogeneity, not allowing conclusions at an acceptable evidence level. Fifth, definitions of complications (e.g., airway trauma, sore throat, and hypoxia) were

not standardized and varied from center to center. Sixth, reported time for tracheal intubation was frequently not reported in the studies included or not normally distributed, prohibiting any statistical analysis. The remaining RCTs showed substantial heterogeneity for tracheal intubation time, not allowing data pooling. Sixth, the protocol for this meta-analysis was registered at PROSPERO after the commencement of the systematic review process. Finally, formal statistical tests did not support the presence of publication bias for any considered outcome, which could have impacted the estimated pooled effect.

Conclusions

Our results suggest that VLSs reduced the risk of difficult intubation and airway trauma while slightly increasing successful intubation at the first attempt among adult patients with a high quality of evidence.

Ethical Approval and Consent to Participate

Not required.

Conflicts of Interest

The authors declare that they have no conflict of interests.

Availability of Data and Materials

Further data and material can be accessed by contacting the corresponding author.

Funding

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Authors' Contributions

MV, GS, PB, CI, PP, CP: study design, data collection, data analysis, writing up the draft of the paper, approved the final version. GPH, SE, LB, NP: data collection, data analysis, writing up the draft of the paper, approved the final version. MA: language editing, grammar check, approval of final version. All authors have read and approved the manuscript.

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