

Routine practice in mechanical ventilation in cardiac surgery in Italy

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Background: Management of mechanical ventilation is a key issue in the prevention of postoperative pulmonary complications (PPCs) and the improvement of surgical outcome. This is especially true in cardiac surgery where the use of the cardiopulmonary bypass (CPB) increases the risk of lung injury. In the last years a growing number of studies have shown that protective ventilation has led to excellent results. However, the literature in this regard is lacking in cardiac surgery and there are no univocal guidelines in this sense. The aim of this survey was to investigate the actual clinical practice about ventilation techniques used in the Italian cardiac surgery centers.

Methods: A questionnaire of 32-item was sent to 69 Italian cardiac surgery centers, 56 of which return a completed form (81.2%). The questionnaire was assembled by three independent researchers and the final version was e-mailed to all members of the SIAARTI (Italian society of anesthesia resuscitation and intensive care medicine) Study Group on Cardiothoracic and Vascular Anesthesia. The answers were collected using a Google Forms sheet. In case of multiple questionnaires returned from the same center (i.e., different physicians from the same center responded) the head of department was asked to give a definite answer. Furthermore, for the 17 centers who reported multiple questionnaires, no large differences were found between the responses of different doctors belonging to the same center (12.3%±4.2% of discordant answers).

Results: Intraoperatively, patients were ventilated with a tidal volume (TV) of 6–8 mL/kg (91.1% of centers), a positive end-expiration pressure of 3–5 cmH₂O (76.8% of centers) and a fraction of inspired oxygen (FiO₂) of 50–80% (60.7% of centers). During the CPB, the "stop ventilation" technique was frequently adopted (73.2%). Before the discharge from the intensive care unit (ICU) non-invasive ventilation (NIV) was never applied in 32.1% of the centers, but it was used in 46.4% of patients with postoperative complications.

Conclusions: This study shows a significant heterogeneity in ventilatory techniques among the Italian centers during CPB, whereas in the other surgical time the majority of the responding centers adopted a protective mechanical ventilation strategy.

Keywords: Mechanical ventilation; cardiopulmonary bypass (CPB); cardiac surgery; survey; pulmonary complications

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Introduction

Management of mechanical ventilation is among the most important skills for a modern anesthesiologist. Securing the airway and providing controlled or assisted mechanical ventilation are fundamental requirements for safe delivery of anesthesia. In recent decades, much progress has been made in protective lung strategy during surgery, including low tidal volume (TV), low plateau and driving pressure, recruitment maneuvers (RMs), and adequate positive endexpiratory pressure (PEEP) (1-3).

This is especially essential in cardiac surgery since many factors can contribute to lung injury, including general anesthesia itself, cardiopulmonary bypass (CPB), blood transfusions, cardiac failure, and diaphragmatic dysfunction—all of which increase the risk of postoperative pulmonary complication (PPC), as defined by Abbott *et al.* (4) in up to 25% of patients after surgery (5-7). Nevertheless, evidence regarding the best way for lung protective ventilation is still lacking.

Based on the assumption that there is no uniformity in the ventilatory management of the cardiac surgery patient in Italian hospitals, we conducted a survey to investigate and understand the current clinical practices in our country, Italy.

Methods

From April 2017 to April 2018, we identified 69 centers performing adult cardiac surgery, and an electronic 32item questionnaire was sent to the 56 centers (81.2%) that accepted our invitation to take part in the survey. The questionnaires were first sent in April 2017, followed by monthly reminders addressed specifically to non-responders until April 2018. The dataset supporting the conclusions of this article is included within the article and Supplementary file 1. Main data graphs are included in http://fp.amegroups. cn/cms/jtd.2019.03.04-1.pdf, http://fp.amegroups.cn/cms/ jtd.2019.03.04-2.pdf.

Our work was endorsed by the Italian Society of Anesthesia, Analgesia, Resuscitation and Intensive Care (SIAARTI). We sent an e-mail invitation to all members of the SIAARTI Study Group on Cardiothoracic and Vascular Anesthesia. The centers performing cardiac surgery in Italy were identified through the Italian Society of Cardiac Surgery (SICCH) website (http://www.sicch.it/), and further information for Cardiothoracic intensive care units (ICUs) were obtained from hospital websites and personal contacts. There were no specific inclusion criteria for the centers. All participating anesthesiologists were informed about the aims of the study. If the chiefs of these ICUs did not respond to our first e-mail, they were personally recontacted by e-mail and invited to participate.

Respondents were asked to indicate one (or more, when necessary) answer to each question using the Google Forms online platform. The survey could not be submitted unless completed since all questions were flagged as mandatory. The respondents were unable to review their answers. All information collected was protected, and no personal contact information was accessible to third parties.

A case report form (CRF) (Supplementary file) consisted of 31 questions regarding both intra- and post-operative issues and was designed to evaluate each participant. When more than one questionnaire was returned by the same hospital (i.e., more than one physician from the same hospital answered the first e-mail), divergences between answers given in the questionnaires were resolved by contacting the head of that department. In this survey, this happened for 17 centers.

The questionnaire sought to investigate the most important aspects of the ventilation setting and use of lung protection strategies in cardiac surgery, both in the intraoperative and postoperative periods. All phases were analyzed, with special attention given to the complex phase of CPB. Three researchers independently developed the items of the survey before the final selection and collection via the questionnaire.

By taking part in the survey, each physician authorized the use of the data recorded in the questionnaire. Informed consent for publication was obtained at the time of participation. The consent manifestation, utilization, and communication of the data collected were performed according to the European General Data Protection Regulation (2016/679, in place as of May 25, 2018). Due to the study design, no ethical approval was required. Specific data regarding individual patients was not collected and therefore remained completely anonymous. This research was carried out in compliance with the Declaration of Helsinki.

Statistical analyses were performed using IBM SPSS Statistics 25.0 (IBM, Armonk, NY, USA) and GraphPad Prism, version 6.01 for Windows (GraphPad Software, San Diego, CA, USA). A descriptive statistical analysis was carried out. Journal of Thoracic Disease, Vol 11, No 4 April 2019



Figure 1 Participating centers. Number of cardiac patients operated per participating center.

Results

A complete list of the answers to the questionnaire is summarized in Supplementary file 1. Further analyzes were made by stratifying the centers by volume and teaching and not-teaching centers; the results of these analyzes are reported respectively in the http://fp.amegroups.cn/ cms/jtd.2019.03.04-1.pdf, http://fp.amegroups.cn/cms/ jtd.2019.03.04-2.pdf.

Center characteristics

The questionnaire was completed and returned by 81.2% of the participant centers, 66.1% of which were non-teaching hospitals, while 33.9% were teaching hospitals. A complete list of the responding centers is available in http:// fp.amegroups.cn/cms/jtd.2019.03.04-3.

High patient flow (>1,000 cardiac surgery procedures per year, with or without CPB) was reported at 7 centers (12.5%). Eight centers (14.3%) reported 750–1,000 procedures per year; 12 centers (21.4%) reported 500–750, 23 (41.1%) centers performed 250–500 procedures per year; and 6 centers (10.7%) performed <250 procedures per year (*Figure 1*).

Type of ventilation in operating room

Low TV ventilation (regardless of the method of ventilation chosen, either pressure-controlled or volume-controlled intermittent positive-pressure ventilation) was used in 91.1% of centers, with TVs of 6 mL/kg (26.8%), 7 mL/kg (33.9%), and 8 mL/kg (30.4%), while a TV of 8-10 mL/kg was used in 8.9% of the centers (Figure 2). The TV was calculated on the ideal body weight in 57% of the centers and on the real body weight in the other 43%. An average PEEP of 3-5 cmH₂O was used in 76.8% of centers, and 5-10 cmH₂O in another 16.1%. Zero PEEP was used in 7.1% of centers. The "best PEEP" was assessed in only 1.8% of the centers intraoperatively (Figure 3). A definition of best *PEEP* was not provided by the questionnaire since there are many methods to assess the best PEEP in clinical practice. The results simply describe the attempt to set a best PEEP value. In 60.7% of the centers, fraction of inspired oxygen (FiO_2) was between 50–80% at the end of surgery. In the other centers, patients were managed with an FiO₂ of less than 50%. We documented a wide range of FiO_2 applied during weaning from CPB: 73.2% of the centers applied an FiO₂ of 50-80%, while 14.3% used an FiO₂ of more than 80%. The 39% of centers use a $FiO_2 < 50\%$ during surgery (Figure 4) while only 12.5% of centers used an FiO₂ of <50% at the weaning from CPB.

Ventilation was generally stopped during CPB (75% of the total centers). Continuous positive airway pressure (CPAP) was applied in 16.1% of the total centers, while only 8.9% did not stop ventilation during CPB (*Figure 5*).

In centers where ventilation was continued during CPB, a CPAP of 5–8 cmH₂O was used in 71.4% of centers, with <5 cmH₂O in the other 21.4% of centers; a CPAP higher than 8 cmH₂O was reported in only 7.1% of centers. In those centers, TV and PEEP values varied from a very low TV of 2–3 mL/kg in 73.3% of the centers to a 3–5 mL/kg TV in 26.7%. A PEEP level of 3–5 cmH₂O was used in 52.6% of the centers, while a PEEP >5 cmH₂O was used in 5.3%. A PEEP of 1–3 cmH₂O was used in 31.6%, with zero PEEP in the remaining 10.5%.

When ventilation was stopped, almost one-third of centers disconnected patients from the anesthesia circuit.

RMs were used at 43 centers (76.8%) and were performed manually at 38 centers (88% of the centers



Figure 2 Tidal volume. Values of tidal volume used before and after cardiopulmonary bypass.



Figure 3 PEEP level. Level of PEEP used to ventilate the patient before and after cardiopulmonary bypass. PEEP, positive end-expiratory pressure.

performing RMs).

Ventilation during patient transport to the ICU

In 12.8% of centers, a fast-track protocol was in use, defined as "extubation carried out in the operating room." When mechanical ventilation was continued in the ICU, manual ventilation was used in 87.5% of centers, while 12.5% used a portable mechanical ventilator.

Type of ventilation in ICU

On arrival in the ICU, the most widely used type of ventilation was synchronized intermittent mandatory

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Figure 4 FiO₂. Values of FiO₂ used at the weaning from cardiopulmonary bypass. FiO₂, inspired fraction of O₂.



Figure 5 Ventilation during CPB. Ventilation method used during cardiopulmonary bypass. MV, mechanical ventilation; CPAP, continuous positive airway pressure; CPB, cardiopulmonary bypass.

ventilation (SIMV) (either pressure-controlled or volumecontrolled SIMV; 41.1% of centers), followed by pressure support (39.3%), bilevel positive airway pressure (BIPAP) (14.3%), volume-controlled mechanical ventilation (VC-CMV) (3.6%), and pressure-controlled mechanical ventilation (PC-CMV) (1.8%).

Once back in the ICU, a TV of 8 mL/kg was used in

32.1% of centers, followed by 7 mL/kg in 32.1%, 6 mL/kg in 25%, and 8–10 mL/kg in 10.7%. PEEP was used in all centers, with values set between 3–5 cmH₂O (66.1% of centers), 6–8 cmH₂O (30.4%), and 8–10 cmH₂O (3.6%). In only 23.2% of the centers was "best PEEP" calculated during the ICU stay. We did not specifically ask the method used to asses a best PEEP, since there is no

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consensus about this important topic among intensive care physicians (8) we only asked if a method was used or the PEEP was simply chosen "clinically", without respiratory mechanics measurements. Respiratory RM were used in 53.6% of centers before extubation. In centers where a fast-track approach was used, a myorelaxant antagonist was administered before extubation in 28.6% of centers, with 65.7% using neostigmine *vs.* 34.3% using sugammadex.

Before the discharge from the intensive care unit noninvasive ventilation (NIV) was never applied in 32.1% of the centers, but it was used in 21.5% of centers for selected patients and in 46.4% of centers only for patients with postoperative complications.

Discussion

The main result of this survey is the lack of a univocal approach among Italian cardiac surgery centers according to the most recent evidence on the best strategy for lung protection in cardiac surgery, a result similar to other surveys (9). However, to our knowledge, we now report this issue in the literature for the first time.

In fact, it is known that most data on protective ventilation come from the experiences of other major surgical centers and mixed ICUs, suggesting the use of low TV, PEEP (10), and, generally speaking, reduction of ventilator-induced lung injury. Moreover, it is also important to calculate the TV on the ideal weight and not on the real one, especially in cases of patients who are severely overweight and obese, where using the increased actual weight to calculate TV targets will overestimate the target TV and expose these patients to harmful volutrauma and barotrauma (11). Two recent meta-analyses showed how protective ventilatory strategies could help reduce PPCs during general anesthesia, as well as possibly shorten the length of a hospital stay (12,13). However, there is still no significant evidence in the literature regarding the most suitable ventilatory strategy to use in cardiac surgery.

Nonetheless, lung damage in cardiac surgery seems to follow the main pathways of barotrauma (high transpulmonary pressure), atelectasis (lack of adequate lung recruitment, particularly if ventilation is stopped during CPB), and inflammation (both ventilation-induced and because of CPB itself) (5,14,15). High plateau pressures and driving pressures (generating high transpulmonary pressures) are likely to damage lung parenchyma in cardiac surgery as well as in general surgery. Moreover, cardiac surgery can present further possible sources of lung injury, such as the inflammatory response induced by CPB (6), complete collapse of lung parenchyma caused by interruption of ventilation during CPB (5,16,17), injury induced by blood transfusion (18), production of proinflammatory cytokines linked to CPB-related myocardial damage, and the production of free radicals following reperfusion of myocardial tissue after CPB (19,20).

In this context, lung-protecting ventilation strategies including the use of low TVs, RMs, adequate FiO₂, avoiding hyperoxia (i.e., absorption atelectasis), and NIV or CPAP during postoperative ICU stay where indicated are thought to play key roles in lung protection. However, there is no high-quality evidence available in the context of cardiac surgery and cardiac ICU. In this way, the lack of consensus in the current practice is not surprising. New research in the field of mechanical ventilation in cardiac surgery is warranted.

FiO2 management

Regarding FiO₂ management, a recent review investigated the effects of oxygen fraction, concluding that moderate hyperoxia (50–80% FiO₂) is potentially beneficial owing to the reduced incidence of surgical-site infections and the absence of demonstrated clinical drawbacks (21). However, conflicting opinions are still present in the literature (22-24). From a pathophysiological point of view, we can say that hemoglobin saturation higher than 100% is not possible, and the fraction of oxygen not bound to hemoglobin carries an insignificant percentage of oxygen delivery at 1 atm. Therefore, it is questionable to keep partial oxygen pressure higher than needed to saturate hemoglobin.

A recent trial compared moderate hyperoxic targets to near-physiological oxygen targets during and after coronary artery bypass surgery, with myocardial damage as a primary end-point (25). The use of a normoxemic strategy did not affect the incidence of myocardial damage, nor did it influence secondary outcomes (such as cardiac index, systemic vascular resistance index, serum creatinine and lactate. A recent meta-analysis based on 12 randomized controlled trials (RCTs) showed the minimal effect of hyperoxia on organ dysfunction, length of hospital stay, and mortality in adult cardiac surgery (26). An ongoing study is currently comparing the increase in serum creatinine in hyperoxygenated patients (FiO₂ of 100%) and those undergoing physiological oxygenation (27).

High oxygen concentrations are associated with the development of absorption atelectasis (28), which can

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lead to the onset of PPC. At present, there is no strong evidence as to how a high FiO_2 might prevent infection in the surgical site and further the development of hyperoxia-induced atelectasis predisposing to postoperative pulmonary infections. There is currently no strong evidence on what might be considered best treatment, with further studies required.

The use of RMs and PEEP could reduce the incidence of atelectasis. Although many studies agree on the usefulness of RMs, a best method has yet to be defined, as does the ideal pressure to reach during the maneuver (28). A recent study carried out during the postoperative period investigated whether an intensive alveolar recruitment strategy yields better results than does a moderate strategy. All patients were ventilated with low-to-moderate TV. The intensive strategy, involving three cycles of lung inflation (60 seconds each), consisting of PEEP of 3 cmH₂O, pressure-controlled ventilation, driving pressure of 15 cmH₂O, respiratory rate of 15/min, inspiratory time of 1.5 seconds, and FiO₂ of 40% was better at preventing PPCs and able to reduce the incidence of severe pulmonary complications (29).

Ventilation during CPB

Although no unequivocal consensus currently exists, there are three options for the management of ventilation during CPB:

- (I) CPAP: various studies used CPAP with pressures between 5–15 cmH₂O and showed different results;
- (II) Mechanical ventilation: low TV frequency ventilation as postoperative oxygenation showed a positive effect on secondary outcomes;
- (III) Lung rest: this seems to be the best option for the surgeon. however, the studies investigated in two systematic reviews showed no significant differences in surgical times compared to experimental arms (27,30).

None of the studies investigated in the two reviews showed damage caused by intra-CPB ventilation, nor by preor post-CPB protective ventilation (6,31,32). However, not every study involved human subjects or cardiac surgery (33).

There is no definitive evidence in the literature showing the superiority of a specific method of ventilation; further studies are required.

Postoperative NIV

NIV can be used both to prevent and to treat PPCs. A

recent study by Olper *et al.* (34) promotes the efficacy of early NIV applications in the cardiac surgery ward, with improved oxygenation in patients who develop hypoxemic acute respiratory failure after discharge from the ICU. It seems that even high-flow nasal oxygen gives results comparable to classic NIV (35).

Limitations of the study

A strong limitation of this study is the differences in clinical practices among anesthesiologists at the same center. When anesthesiologists at the same center gave significantly different answers, the head of the center was contacted to clarify the data.

As a general limitation of the survey, we cannot verify whether what a single physician declares is truly correct and reflects the current practice in his or her center. Further, if a survey uses different words or phrases from those in a particular clinical practice, the results of that survey could have a different story to tell. Moreover, we did not receive answers from all the centers in Italy; therefore, we cannot assume our data to be conclusive. In the non-responding centers, there may be differences in clinical practice from the responses we were able to collect.

Different physicians can have varying opinions regarding management of the same patient based on their personal experience. However, our questionnaire was designed to investigate procedures that generally respected the established protocol of each specific center.

Conclusions

This survey describes the current state of Italian ventilation management during cardiac surgery and shows that there is still some heterogeneity among ventilation settings, in particular, during CPB (although protective ventilation in other surgeries is widespread).

Focusing on current scientific evidence, standards of care need to be established concerning ventilation of patients during cardiac surgery at Italian centers. Further research is needed to investigate the methods of protective ventilation, as these techniques have already produced important results in other contexts.

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Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

Ethical Statement: This research was carried out in compliance with the Declaration of Helsinki. Informed consent for publication was obtained at the time of participation.

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A complete list of the answers to the questionnaire

Section A: general	data								
0. Type of center									
Nonteaching	66%		Teaching		34%				
1. Number of cardiac surgery patients operated for year:									
<300	11%	250–500	41%	500-750	21%	750–100	14%	>1,000	13%
Section B. intraope	erative ventils	ation							
2. With which Tida	al Volume do	you ventilate	your patients?						
6 mL/kg	27	7%	7 mL/kg	34	%	8 mL/kg	30%	8–10 mL/kg	9%
3. Is the tidal volu	me generally	/ calculated o	n the ideal weig	ght or on the	actual	weight?			-
Ideal weight	57	7%	Real weight	43'	%	-			
4. With what level	of positive e	end-expiratory	y pressure (PEE	P) do you ve	ntilate	your patients on ave	rage?		
Zero PEEP	7	%	3–5 cmH2O	77	%	5–10 cmH2O	16%	>10 cmH ₂ C	0%
5. Do you use any	method to	calculate a be	est PEEP during	surgery?					
Yes	2	%	No	98	%				
6. If yes, which or	e?								
One answer: cor	npliance								
7. Which inspired	fraction of C)2 (FiO2) gener	ally do you use	during surge	ery?				
<50%	39	9%	50-80%	61	%	>80%	0%		
8. Which FiO2 do	∕ou use at th	ne weaning fro	om cardiopulmo	onary bypass	?				
<50%	13	3%	50-80%	73	%	>80%	14%		
9. Which ventilation	on method u	sed during th	e cardiopulmon	ary bypass?					
Stop ventilation	75	5% Con	tinuation mecha	anical 9%	6	Continuous positive	16%	Other	0%
			ventilation			airway pressure			
10. If you continue	e ventilation	during cardio	pulmonary byp	ass, with whi	ch tida	l volume?			
2–3 mL/kg	73	3%	3–5 mL/kg	27	%	>5 mL/kg	0%		
11. If you continue	e ventilation	during cardio	pulmonary byp	ass, with whi	ch PEE	P?			
Zero PEEP	11	1%	1–3 cmH ₂ O	32	%	3–5 cmH ₂ O	53%	>5 cmH₂O	4%
12. If you use CPA	AP during ca	rdiopulmonar	y bypass, what	pressure leve	el do yo	ou set?			
<5 cmH₂O	21	1%	5–8 cmH2O	72	%	>8 cmH₂O	7%		
13. If you switch c	off the ventila	ator during ca	rdiopulmonary	bypass, do y	ou disc	connect the patient fr	om the circ	uit?	
Yes	28	3%	No	72	%				
14. Do you perfor	m recruitmer	nt maneuvers	during surgery	?					
Yes	77	7%	No	23	%				
15. If you perform	recruitment	maneuvers, v	with which mod	le?					
Manual	83	3%	In	creasing volu	imes/pi	ressures gradually fro	om the vent	ilator 17%	
16. Do you give co	ortisone-bas	ed drugs rout	tinely during su	rgery?					
Yes	14	1%	No	86	%				
17. Do you use ar	y other syst	em for lung p	rotection during	g surgery?					
Yes	2	%	No	98	%				
18. If yes, which c	ne?								
One answer				Ha	logena	ted anaesthetics			

Section C: intensive care unit transportation

Yes46%No54%20. If you apply a fast track protocol, do you extubate the patients in the operating theater?	19. Do you apply a fast tra	ack protocol in your center?		
20. If you apply a fast track protocol, do you extubate the patients in the operating theater?	Yes	46%	No	54%
	20. If you apply a fast trac	k protocol, do you extubate the patie	ents in the operating theater?	
Yes 13% No 87%	Yes	13%	No	87%
21. How do you ventilate patients during transport?				
Manually87%Portable ventilator13%	Manually	87%	Portable ventilator	13%

Section D: postoperative mechanical ventilation during ICU stay

22. How do you ventilate patier	its during Inte	nsive Care Unit st	tay after the	neuromuscular blockingag	gent effec	t has finished?	
Synchronized intermitted mandatory ventilation	41%	Pressure support	39%	Bilevel positive airway pressure	14%	Other	6%
23. Which tidal volume target de	o you have in	post-operative int	ensive care	unit stay?			
6 mL/kg	25%	7 mL/kg	32%	8 mL/kg	32%	8–10 mL/kg	11%
24. Do you use PEEP during po	st-operative ir	ntensive care unit	stay?				
Yes	100%	No	0%				
25. If you use PEEP in the post-	-operative, wh	at value do you g	enerally use	ə?			
3–5 cmH₂O	66%	6–8 cmH₂O	30%	8–10 cmH2O	4%	>10 cmH ₂ O	0%
26. Do you use a method to cal	culate the bes	t PEEP during Int	ensive Care	Unit stay?			
Yes	23%	No	77%				
27. Do you routinely carry out re	ecruitment ma	neuvers before e	xtubation?				
Yes	54%	No	46%				
28. Do you routinely perform me	easurements o	of respiratory mec	hanics befo	re extubation?			
Yes	23%	No	77%				
29. Do you antagonize the neur	omuscular blo	cking agent befor	re extubatio	n?			
Yes	0%	No	100%				
30. In the case of fast track (if c	arried out), do	you antagonize t	he neuromu	iscular blocking agent befo	re extuba	tion?	
Yes	29%	No	71%				
31. What drugs do you routinely	vuse to antag	onize the effects o	of Neuromu	scular blocking agent?			
Neostigmina	66%	Sugammadex	34%				
32. Do you use non-invasive ve	entilation after	discharge from In	tensive Car	e Unit?			
Yes, routinely	0% Y	es, in selected patients	22%	Yes, only in complicated 46% patients	No	32%	