Safety and Applicability of Continuous Retrograde Cardioplegia in Minimally Invasive Aortic Valve Replacement: New Approaches

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Purpose: To discuss minimally invasive cardiac surgery aortic valve replacement (MICS-AVR) approach via anterior thoracotomy using continuous retrograde cardioplegia. Continuous retrograde cardioplegia facilitates excellent continuous homogeneous cooling of the heart during cardiac arrest.

Methods: We performed AVR using the proposed method in nine patients between June 2018 and September 2019. The median age of the patients was 73 (range: 43–84) years. The pleural space was entered via anterior thoracotomy. After opening of the right atrium, a retrograde cardioplegic cannula was inserted into the coronary sinus with a purse-string suture. Continuous cold blood retrograde cardioplegia was initiated at 700 mL/h.

Results: Extubation in the operating room was performed in five (56%) patients. No new decreased function of the left and right ventricles was observed in intraoperative transesophageal echocardiography or transthoracic echocardiogram.

Conclusion: MICA-AVR through continuous retrograde cardioplegia is a safe technique.

Keywords: minimally invasive cardiac surgery aortic valve replacement, superior vena cava, inferior vena cava

Introduction

There are some disadvantages of minimally invasive cardiac surgery aortic valve replacement (MICS-AVR) through thoracotomy. For instance, it leads to complications, such as selective cardioplegic perfusion and difficulty in tying down the biological valve to the aortic annulus. This study aimed to discuss MICS-AVR via anterior thoracotomy through continuous retrograde cardioplegia. Continuous retrograde cardioplegia facilitates excellent continuous homogeneous cooling of the heart during cardiac arrest.1

Materials and Methods

We performed MICS-AVR using the proposed method in nine patients between June 2018 and September 2019. The median age of the patients was 73 (range: 43–84) years. We obtained IRB approval (number of IRB approval: No: 2020-019) on June 9, 2020.

Operative procedure

The right upper part of the patient’s body was slightly bolstered, and both the arms were placed along the side...
of the body. Under general anesthesia with transesophageal echocardiography monitoring and one-lung ventilation, the pleural space was entered via the third intercostal space by cutting the fourth costal cartilage. If ascending aortic cannulation or replacement of the ascending aorta was required, the third costal cartilage was cut. In female patients, we could enter the third intercostal space via the submammary sulcus using epifascial exfoliate.

The heart and aorta were moved closer to the right anterior chest wall by pulling up of the pericardium, which is similar to the Yamazaki’s Stonehenge technique. Patients whose ascending aorta was located to the left of the midline (Fig. 1) and those with a relatively short distance between the sternum and vertebra could be treated without any difficulties. The right lung was separated from the operative field using the pericardium and traction strings thereafter, and one-lung ventilation was then discontinued.

Cardiopulmonary bypass was performed via femoral/ascending aorta cannulation with 20Fr cannula (femoral; Medtronic, Mineapolis, MN, USA, ascending aorta; Mera) and femoral drainage with 25Fr cannula. Superior vena cava (SVC) drainage with 20 or 22Fr flexible L-shape cannula was established via incision. The blood temperature was cooled to 34°C.

Tapes were placed around the SVC and the inferior vena cava (IVC). A vent tube was inserted via the right upper pulmonary vein. Antegrade cardioplegic cannula was placed in the ascending aorta.

The ascending aorta was clamped, and cardiac arrest was induced with cold antegrade cardioplegia (9°C).

After opening of the right atrium, a retrograde cardioplegic cannula was inserted into the coronary sinus with a

![Fig. 1](image_url)

**Fig. 1** (A) Even in cases where the aorta was located slightly to the left of the sternum, the operative field was still good with anterior intercostal thoracotomy and pericardial traction. (B) Same patient, on three-dimensional computed tomography scan, the ascending aorta was not observed from the right side of the sternum. (C) and (D) Same patient, we could obtain a good operative field with suture traction and twisting of the upper body.
A balloon catheter was utilized for retrograde cardioplegic cannula and was placed as shallow as possible in the coronary sinus using purse-string suture. Continuous cold blood retrograde cardioplegia was initiated in 700 mL/h. After opening the aorta, it was found that there was sufficient backflow from the coronary ostia and it was red, not black. If the backflow was inadequate, the flow rate of retrograde cardioplegic fluid was increased.

The composition of undiluted cardioplegia is described below: sodium ion, 132 mEq/L; potassium ion, 52 mEq/L; calcium ion, 2.2 mEq/L; HCO₃⁻, 50 mEq/L (theoretical values); and pH, 7.8 (the actual measurement value). In continuous retrograde cardioplegia, the ratio of undiluted cardioplegia to blood was 1:5. The infusion pressure of retrograde cardioplegia was <40 mmHg. The solution temperature was 9°C.

AVR was performed in a similar manner. During aortic valve replacement, antegrade cardioplegic cannula was set aside. After providing warm cardioplegia (38°C), the aortic clamp was removed.

The resected costal cartilage was fixed with a monofilament absorbent suture.

## Results

Extubation in the operating room was performed in five (56%) patients (Table 1). None of the cases had to wait specifically for the recovery of cardiac function until weaning from cardiopulmonary bypass, and no case required defibrillation after aortic declamping. None of the cases required intra-aortic balloon pumping or percutaneous cardiopulmonary support. No new decreased function of left and right ventricles was observed in intraoperative transesophageal echography or postoperative transesophageal echocardiogram or pressure data at the intensive care unit entry (Table 2). One patient required return cardiac arrest due to perivalvular leakage. As the number of cases increased, the number of muscle fiber cuts decreased due to the reduction in the wound and the incision along the muscle fibers, and postoperative creatine kinase (CK) rise was suppressed.

## Discussion

There is no report of MICS-AVR using continuous retrograde cardioplegia. Continuous retrograde cardioplegia is the rational approach in MICS-AVR. There are two advantages of retrograde cardioplegia. One, complications due to selective cardioplegia are prevented, and two, it is useful for de-airing.

There are two advantages of continuous cardioplegia. One, it is excellent in terms of myocardial protection. For example, Louagie et al. reported that continuous

| No. | Age (y) | Sex | AR/AS | Cannula- tion | OP time (min) | CPB (min) | Ax (min) | CK-MB (ng/mL) | CK (ng/mL) | Transfusion | Extubation | Discharge (days after operation) |
|-----|---------|-----|-------|--------------|---------------|-----------|----------|---------------|------------|-------------|------------|----------------------------------|
| 1   | 71      | Female | AS | FA | 370 | 211 | 144 | 26 | 426 | + | ICU | 8 |
| 2   | 43      | Male | AR | FA | 305 | 183 | 93 | 65 | 3129 | - | Operative theater | 14 |
| 3   | 77      | Male | AS | AscAo | 365 | 228 | 135 | 57 | 2568 | - | Operative theater | 11 |
| 4   | 78      | Female | AS | AscAo | 525 | 332 | 196 | 67 | 2088 | + | ICU | 23 |
| 5   | 71      | Female | AS | FA | 273 | 175 | 111 | 55 | 377 | - | Operative theater | 11 |
| 6   | 74      | Male | AR | FA | 226 | 125 | 69 | 35 | 369 | - | Operative theater | 10 |
| 7   | 84      | Female | AS | FA | 284 | 188 | 117 | 30 | 336 | + | ICU | 25 |
| 8   | 49      | Male | AR | FA | 294 | 201 | 108 | 36 | 1410 | - | ICU | 8 |
| 9   | 59      | Male | AR | FA | 229 | 141 | 78 | 36 | 383 | - | Operative theater | 8 |

No. reflects the order of operation.

*Return cardiac arrest was required due to perivalvular leakage.

AR: aortic regurgitation; AS: aortic stenosis; AscAo: ascending aorta; Ax: cardiac arrest time; CK: creatine kinase immediately after finishing the operation; CPB: cardiopulmonary bypass time; FA: femoral artery; ICU: intensive care unit; OP time: operation time.
cold blood cardioplegia results in fewer cardiac events, better left ventricular performance, and a marked improvement of the right ventricular function when compared with similar solutions of blood cardioplegia administered intermittently, independent of their means of delivery.

In Louagie's study, the infusion speed of continuous retrograde cardioplegia was 50 mL/min; however, our target amount of continuous cardioplegia showed approximately 700 mL/h, which is the same as the additional amount of intermittent cardioplegia per hour. It is presumed to be sufficient for cooled myocardium. We believe that excessive cardioplegia may cause myocardial edema. According to Higami et al., the amount of continuous retrograde cold cardioplegia perfused per left ventricular mass weight (LVMW) was significantly correlated with mitochondrial damage. This study is a model performed by a method of cardioplegia similar to our study, wherein the right ventricular myocardium was biopsied and mitochondrial damage was observed using an electron microscope. In cases where cold cardioplegia was administered at 5 mL/100 g LVMW/hour, mitochondria with irreversible damage were not observed.

The average left ventricular myocardial weight in the Japanese population is estimated to be 133 ± 28 g for men and 105 ± 22 g for women, according to Kato et al. [6]. The average amount of cold cardioplegia used in our study was 5 mL/100 g LVMW/hour, which is the same as the additional amount of intermittent cardioplegia per hour. This study is a model performed by a method of cardioplegia similar to Louagie's study, the infusion speed of continuous retrograde cardioplegia was 50 mL/min; however, our target amount of continuous cardioplegia showed approximately 700 mL/h, which is the same as the additional amount of intermittent cardioplegia per hour. It is presumed to be sufficient for cooled myocardium.

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insertion of retrograde cardioplegia cannula into the coronary sinus via the right atriotomy causes trouble-free cardioplegia.

The following are the advantages of this type of surgery: this approach is similar to or even more convenient than partial sternotomy; it can be applied easily to a patient with aortic regurgitation who requires retrograde cardioplegia; no selective cardioplegia, which is a complicated procedure, is required; continuous perfusion can prevent myocardial ischemia; and cannulation of the ascending aorta can be selected if femoral cannulation is not possible.

Meanwhile, the disadvantages of this surgery are as follows: right atrium incision and cutting of the fourth costal cartilage are required, and hemofiltration is often performed in cardiopulmonary bypass for potassium control because the volume of cardioplegic solution is high.

Conclusion
MICS-AVR via anterior thoracotomy using continuous retrograde cardioplegia is a safe technique.

Disclosure Statement
None.

References

1) Bezon E, Choplain JN, Khalifa AA, et al. Continuous retrograde blood cardioplegia ensures prolonged aortic cross-clamping time without increasing the operative risk. Interact Cardiovasc Thorac Surg 2006; 5: 403–7.
2) Yamazaki M, Kin H, Kitamoto S, et al. Efficacy of the Stonehenge technique for minimally invasive aortic valve replacement via right infraaxillary thoracotomy. Ann Thorac Surg 2017; 23: 45–8.
3) van Putte BP, Vink A, De Bruin PC, et al. Selective antegrade cardioplegic perfusion complicated by left main stem dissection. J Cardiovasc Surg 2007; 48: 247–8.
4) Yates JD, Kirsh MM, Sodeman TM, et al. Coronary ostial stenosis, a complication of aortic valve replacement. Circulation 1974; 49: 530–4.
5) Louagie YA, Gonzalez E, Jamart J, et al. Assessment of continuous cold blood cardioplegia in coronary artery bypass grafting. Ann Thorac Surg 1997; 63: 689–96.
6) Higami T, Asada T. Protective effect of the retrogradely perfused cold blood cardioplegia to the right ventricular myocardium. Electron microscopical study of mitochondria. Jpn J Cardiovasc Surg 1989; 19: 527–9.
7) Daimon M, Watanabe H, Abe Y, et al. Normal values of echocardiographic parameters in relation to age in a healthy Japanese population: the JAMP study. Circulation 2008; 72: 1859–66.