Evolution of the management approach for pulmonary atresia with intact ventricular septum

Y P Mi, A K T Chau, C S W Chiu, T C Yung, K S Lun, Y F Cheung

Objective: To review the evolution of the management approach for pulmonary atresia with intact ventricular septum (PAIVS) in the past two decades and to assess its impact on patient outcomes.

Design and patients: Retrospective review of the management and outcomes of 94 patients (55 male patients) with PAIVS diagnosed between July 1980 and August 2003.

Settings: Tertiary paediatric cardiac centre.

Results: Seven patients died before interventions. Of the remaining 87 patients who underwent intervention at a median age of 9 days (from 1 day to 2 years), 12 had right ventricular outflow tract reconstruction (RVOTR), 42 had closed pulmonary valvotomy (CPV), and 15 had laser assisted valvotomy with balloon valvoplasty. A systemic–pulmonary shunt was inserted in 18 patients, six of whom had subsequent RVOTR (n = 4) or laser assisted valvotomy (n = 2). Since 1990, catheter intervention accounted for 38% (17 of 45) of the right ventricular outflow procedures. The mean (SEM) freedom from reintervention was 93 (7)%, 71 (12)%, and 57 (13)% after RVOTR, 75 (7)%, 40 (8)%, and 14 (6)% after CPV, and 54 (13)%, 24 (12)%, and 16 (10)% after laser assisted valvotomy at one month, six months, and one year, respectively (RVOTR versus CPV, p < 0.001; RVOTR versus laser assisted valvotomy, p = 0.001). Low cardiac output syndrome was significantly less common after catheter intervention than after RVOTR (0% v 44%, p = 0.003) or CPV (0% v 29%, p = 0.01). The overall mean (SEM) survival was 77 (5)% and 70 (5)% at one and five years, respectively, and the overall mortality was 33% (29 of 87).

There were no significant differences in survival between the three groups.

Conclusions: Multiple interventions are often required in the treatment algorithm of PAIVS. The shift towards increased use of the transcatheter approach has reduced the occurrence of postprocedural low cardiac output syndrome.

METHODS

Patients and data collection
We reviewed the clinical records of 94 patients (55 male patients) with PAIVS diagnosed between July 1980 and August 2003. The cardiac diagnosis was based primarily on two dimensional and Doppler echocardiography, although angiography had been the mainstay of diagnosis in the earlier years.

The following data were retrieved from the clinical records: demographic data including age, weight, and year at presentation; sex; the initial intervention offered, complications including low cardiac output syndrome and the outcome; the need for and type of subsequent additional interventions; for patients who died, the age at death and cause of death; and for survivors, their further management, follow up duration, and outcomes. Low cardiac output syndrome was defined as the presence of clinical signs and symptoms of the syndrome, as tachycardia, oliguria, cold extremities, or cardiac arrest, with or without a ≥ 30% difference in arterial–mixed venous oxygen saturation or metabolic acidosis.17

To determine the morphology of the RV, type of atresia (muscular versus membranous), degree of tricuspid regurgitation, and coronary artery abnormalities, we reviewed all

Abbreviations: CPV, right ventricular outflow tract reconstruction; PAIVS, pulmonary atresia with intact ventricular septum; RV, right ventricular; RVOTR, right ventricular outflow tract reconstruction
the echocardiographic and angiographic recordings at presentation. Morphological description of the RV was based on the tripartite classification described by Bull and colleagues and degree of RV hypoplasia was assessed subjectively. Additionally, the tricuspid valve annulus dimension was measured at end diastole in the echocardiographic apical four chamber view and expressed as a z score from normograms based on postmortem data and contemporary echocardiographic data. The degree of tricuspid regurgitation was estimated as absent, mild, moderate, or severe either angiographically or echocardiographically. Furthermore, the presence of Ebstein’s malformation of the tricuspid valve was noted. The coronary artery anatomy was studied from the RV and aortic root angiograms and the presence of RV to coronary artery fistulae was noted. An RV dependent coronary circulation was defined by the presence of coronary arterial interruption, stenosis of one or more of the major epicardial arteries, or absent aortocoronary connection.

Surgical and transcatheter interventions
Ductal patency was maintained by prostaglandin E2 before the surgical or transcatheter procedure. The management strategy for PAIVS has evolved over the past two decades in our institution. Between July 1980 and October 1991, 12 patients underwent RVOTR as the initial procedure. Between December 1984 and December 1996, transventricular closed pulmonary valvotomy (CPV), performed in 42 patients, had become the mainstay of initial management as reported by our group previously. Since its introduction in January 1996, transcatheter laser assisted valvotomy with balloon pulmonary valvoplasty had been the initial procedure in 15 patients who did not have infundibular atresia and RV dependent coronary circulation. In patients with severe RV hypoplasia or RV dependent coronary circulation and in those born preterm or having a low birth weight, the initial palliative management was a systemic-pulmonary arterial shunt.

After successful surgical or transcatheter establishment of ventriculopulmonary continuity, prostaglandin infusion was discontinued. If severe systemic oxygen desaturation (70%) occurred, prostaglandin infusion was restarted. Systemic–pulmonary arterial shunt was inserted in patients who remained prostaglandin dependent for three weeks or more and earlier in those who developed severe hypoxaemia despite prostaglandin infusion. On the other hand, if the patient showed evidence of progressive heart failure, ductal ligation was performed.

Follow up
All of the survivors had serial echocardiographic assessment to monitor RV growth, severity of residual RV outflow obstruction, and degree of tricuspid and pulmonary regurgitation. The severity of pulmonary regurgitation was graded as the ratio of the colour jet diameter to pulmonary annulus in the parasternal short axis at the level of the RV outflow tract. For patients with significant RV outflow obstruction (a systolic pressure gradient > 50 mm Hg), balloon valvoplasty or RVOTR was performed as required. For those with persistent systemic oxygen desaturation (< 90%) despite the absence of significant RV outflow obstruction, device closure of the atrial communication would be performed if a trial of balloon occlusion was tolerated. A biventricular circulation

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**Figure 1** Flowchart showing interventions and outcomes in patients undergoing primary right ventricular outflow tract reconstruction (RVOTR). ASD, atrial septal defect; LPA, left pulmonary artery; PBPV, percutaneous balloon pulmonary valvoplasty; PDA, patent ductal artery; SP, systemic–pulmonary.

**Figure 2** Flowchart showing interventions and outcomes in patients undergoing primary closed pulmonary valvotomy (CPV).
was considered to have been achieved if there was no systemic oxygen desaturation together with the absence of right to left interatrial shunt as documented by colour flow mapping echocardiographically.

**Statistical analysis**
Results are expressed as mean (SD) or median (range) unless otherwise stated. Differences in demographic data, RV morphological variables, and types of initial interventions between survivors and non-survivors were compared by unpaired Student’s *t* test, Mann-Whitney U test, and Fisher’s exact test where appropriate. Survival of patients and freedom from the need for multiple interventions were analysed by Kaplan-Meier actuarial survival analysis and groups were compared by log rank test. A probability value of *p* ≤ 0.05 was considered significant. All statistical analyses were performed with SPSS version 10.0 (SPSS, Inc, Chicago, Illinois, USA).

**RESULTS**

**Patients**
All but two of the 94 patients were ethnic Chinese. The median age at presentation was 1.5 days (range 1 day to 2 years) and the median weight was 3.2 kg (range 1.4 to 7.2).

Three patients had associated syndrome abnormalities, namely Down’s syndrome, Noonan’s syndrome, and Turner’s syndrome. Balloon atrioseptostomy was performed in 15 patients. The atresia was membranous in 88 (93.6%) and muscular in six (6.4%) patients. Four (4.3%) patients had Ebstein’s malformation of the tricuspid valve. Although 20 (21.3%) had RV to coronary artery fistulae, only one (1.1%) patient had RV dependent coronary circulation diagnosed.

**Surgical and transcatheter management**
Seven patients died before any possible interventions. Of the remaining 87 patients who underwent interventions at a median age of 9 days (range 1 day to 2 years), 12 (14%) had RVOTR, 42 (48%) had CPV, 15 (17%) had laser assisted pulmonary valvotomy with balloon valvoplasty, and 18 (21%) had insertion of a modified Blalock-Taussig shunt as the initial intervention. Of the 18 patients with shunt insertion, six eventually had their RV outflow opened by either RVOTR (*n* = 4) or laser assisted valvotomy (*n* = 2). Figures 1 to 4 summarise the various interventions performed and the outcomes.

Before 1990, the RV outflow was rendered patent by surgery, RVOTR in 10 and CPV in 20 patients. On the other hand, of the 45 patients undergoing RV outflow procedures

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**Figure 3** Flowchart showing interventions and outcomes in patients undergoing primary laser assisted valvotomy and balloon valvoplasty.

**Figure 4** Flowchart showing interventions and outcomes in patients undergoing primary systemic–pulmonary (SP) shunt.
after 1990, 17 (38%) had transcatheter interventions, 22 had CPV, and six had RVOTR (p < 0.001).

Regardless of the initial intervention for establishment of RV–pulmonary continuity, the need for one or more additional interventions was common. Overall, 52 of the 75 (69%) patients who had their RV outflow opened required additional interventions. The second interventions were balloon pulmonary valvoplasty in 22 patients, systemic–pulmonary shunt insertion with (n = 6) or without ligation (n = 8) of arterial duct in 14, RVOTR in nine, isolated pulmonary shunt insertion with (n = 6) or without ligation of arterial duct in five, device occlusion of atrial septal defect in five, RVOTR in nine, isolated pulmonary shunt insertion with (n = 6) or without ligation of arterial duct in five, device occlusion of atrial septal defect in one, and balloon angioplasty for left pulmonary artery stenosis in one. Twenty four patients required three interventions and eight patients required more than three interventions.

The mean (SEM) freedom from reintervention was 93 (7)%; 71 (12)% and 57 (13)% after RVOTR, 75 (7)%; 40 (8)% and 56 (6)% after CPV, and 54 (13)%; 24 (12)% and 16 (10)% after laser assisted valvotomy at one month, six months, and one year, respectively (fig 5). The need for reintervention was significantly less after RVOTR than after both CPV (p < 0.001) and laser assisted valvotomy (p = 0.001). However, no significant difference was found between the surgical and laser assisted valvotomy treated patients (p = 0.88).

Complications and mortality

Low cardiac output syndrome was the main complication after RVOTR, occurring in 44% (seven of 16) of patients. Of the 42 patients who had CPV, 12 (29%) developed low cardiac output syndrome, four (10%) required peritoneal dialysis for oliguric renal failure, two (5%) had sepsis, and one (2%) had the complication of subcapsular haematoma of the liver. Among the 17 patients undergoing catheter interventions, sepsis occurred in one (6%) and perforation of RV outflow in another (6%). Low cardiac output syndrome was hence significantly less common after catheter intervention than after RVOTR (p = 0.003) or CPV (p = 0.01).

The overall mortality was 33% (29 of 87). Table 1 summarises the causes of early and late deaths. Early postprocedural mortality (death within one month of intervention) was 31% (five of 16) after RVOTR, 19% (eight of 42) after CPV, and 12% (two of 17) after laser assisted valvotomy. The early mortality after palliative shunt operation was 6% (one of 18). Univariate analyses found no significant differences in demographic data, RV morphological variables, type, and the year of the first intervention between survivors and non-survivors (table 2).

Survival and outcomes

The overall mean (SEM) survival of the 87 patients undergoing interventions was 85 (4)%; 77 (5)% and 70 (5)% at one month, one year, and five years, respectively. With regard to individual procedures, the survival was 81 (10)%; 62 (12)% and 56 (12)% after RVOTR, 81 (6)%; 76 (7)% and 71 (7)% after CPV, and 88 (8)%; 82 (9)% and 82 (9)% after laser assisted valvotomy at one month, one year, and five years, respectively (fig 6). Although survival after primary CPV and laser assisted valvotomy appeared to be better than that after RVOTR, the difference was not significant (p = 0.43 and p = 0.22, respectively).

The 58 survivors were followed up for 11.4 (5.6) years (range 3 months to 23 years). One patient who had CPV and a systemic–pulmonary shunt was lost to follow up after immigration. Of the remaining 57 survivors, 39 (68%) patients achieved a biventricular circulation, seven (12%) had a univentricular repair, and two (3.5%) had a one and a half ventricular repair. Six patients were considered likely candidates for biventricular repair, pending either device occlusion of atrial septal defect or RVOTR. One was considered to be a candidate for one and a half ventricular repair. The remaining two patients were managed conservatively after the initial palliative systemic–pulmonary shunt operation; one of them had hypoplastic pulmonary vasculature and the other had pulmonary hypertension and severe RV hypoplasia.

At the latest follow up, 43 (75%) patients were pink in room air. Of the 39 patients who had completed the biventricular repair, the degree of pulmonary regurgitation was found to be mild in 24, moderate in 13, and severe in two patients echocardiographically. Procedures pending were device occlusion of atrial septal defect in five, RVOTR in one, and left pulmonary artery stenting in one patient.

DISCUSSION

The present clinical series from a single institution illustrates the evolution of management strategies over the past two decades for the challenging condition of PAIVS. In patients with a diminutive RV and RV dependent coronary circulation, the need for reintervention was significantly less after RVOTR than after both CPV (p < 0.001) and laser assisted valvotomy (p = 0.001). However, no significant difference was found between the surgical and laser assisted valvotomy treated patients (p = 0.88).
the management is palliative systemic–pulmonary arterial shunt insertion and subsequent univentricular repair. In those with anatomically suitable RVs, laser assisted valvotomy with balloon valvoplasty has become the procedure of choice in our institution. However, regardless of the initial procedure, multiple interventions are often required in the subsequent treatment algorithm, especially for patients after CPV and laser assisted pulmonary valvotomy with balloon dilatation. Nonetheless, the risk of developing low cardiac output syndrome is significantly less after catheter intervention than after either RVOTR or CPV. Although survival of patients after RVOTR appeared worse than survival after CPV or primary catheter intervention, the difference was not significant.

Survival of patients with PAIVS has been relatively poor until recently. In the multi-institutional study reported by Hanley and colleagues\(^{18}\) that constitutes the largest cohort of surgical patients to date, the overall survival was 64% at four years. The 56% and 71% five year survival of our patients after RVOTR and CPV, respectively, concur with their findings\(^{18}\) and those reported previously.\(^{11,23}\) While transannular patch opening of the RV outflow reduces the chance of residual outflow obstruction, the possibility of acute RV failure, especially in the presence of significant RV hypoplasia (tricuspid valve z value\(^{2}\)), remains an issue of concern.\(^{18,23}\) However, using a selective approach based on the RV growth and coronary artery anatomy, Jahangiri and colleagues\(^{9}\) had an excellent surgical result of a zero mortality in 25 patients after RVOTR with or without concomitant systemic–pulmonary arterial shunt insertion.

Notwithstanding the significantly improved survival after RVOTR, CPV has its advocates.\(^{10,20,22}\) Surgical pulmonary valvotomy is a relatively simple procedure that can be performed by the transventricular\(^{20}\) or transarterial\(^{10}\) route without the need for cardiopulmonary bypass, and it may produce less RV damage and pulmonary regurgitation.\(^{22}\) Furthermore, in the presence of RV to coronary communications, it has been suggested that stepwise reduction of RV pressure by initial survival valvotomy and later balloon valvoplasty may potentially avoid compromising myocardial perfusion secondary to RV steal, as the RV pressure usually remains high after the initial valvotomy.\(^{19}\) However, as our data show (fig 2), additional procedures are often required after CPV. In the early postoperative period, 19% (8/42) of patients required a systemic–pulmonary artery shunt after CPV but in the intermediate and long terms, 64% (27/42) required further relief of residual outflow obstruction by either balloon valvoplasty or RVOTR.

The optimal surgical approach to relieve RV outflow obstruction hence remains controversial. Additionally, the need for concomitant systemic–pulmonary arterial shunt insertion as part of the initial surgery is controversial.

### Table 2

<table>
<thead>
<tr>
<th></th>
<th>Survivors (n = 58)</th>
<th>Non-survivors (n = 29)</th>
<th>p Value</th>
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<td>Age at 1st procedure (days)</td>
<td>8 (1 to 307)</td>
<td>9 (1 to 757)</td>
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<td>Weight at 1st procedure (kg)</td>
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<td>Sex (male/female)</td>
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<td>17/12</td>
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<tr>
<td>RV hypoplasia</td>
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<td>Mild</td>
<td>19</td>
<td>9</td>
<td>0.37</td>
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<tr>
<td>Moderate</td>
<td>15</td>
<td>5</td>
<td></td>
</tr>
<tr>
<td>Severe</td>
<td>24</td>
<td>15</td>
<td></td>
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<tr>
<td>RV morphology</td>
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<td>Tricuspid regurgitation</td>
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<tr>
<td>Nil or mild</td>
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<td>8</td>
<td>0.51</td>
</tr>
<tr>
<td>Moderate</td>
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<td>6</td>
<td></td>
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<tr>
<td>Severe</td>
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<td>15</td>
<td></td>
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<tr>
<td>Ebstein’s abnormality (yes/no)</td>
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<td>2/27</td>
<td>0.60</td>
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<tr>
<td>Tricuspid valve dimension</td>
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<td></td>
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<tr>
<td>Annulus size (mm)</td>
<td>10.4 (2.9)</td>
<td>9.8 (3.4)</td>
<td>0.49</td>
</tr>
<tr>
<td>Annulus z score(^{2})</td>
<td>−1.2 (1.5)</td>
<td>−1.5 (2.4)</td>
<td>0.52</td>
</tr>
<tr>
<td>Annulus z score(^{2})</td>
<td>−3.8 (2.5)</td>
<td>−3.9 (2.7)</td>
<td>0.85</td>
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<td>Type of primary procedure</td>
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<td>RVOTR</td>
<td>7</td>
<td>5</td>
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<td>CPV</td>
<td>30</td>
<td>12</td>
<td>0.24</td>
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<td>Laser assisted valvotomy</td>
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<tr>
<td>SP shunt</td>
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<td>9</td>
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<tr>
<td>Year of intervention (before/after 1990)</td>
<td>26/32</td>
<td>9/20</td>
<td>0.25</td>
</tr>
</tbody>
</table>

Data are median (range), mean (SD) or numbers.

*Calculated based on postmortem\(^{18}\) data; \(^{2}\)based on echocardiographic data.\(^{19}\)

RV, right ventricular.

### Figure 6

Kaplan-Meier survival estimates for patients after primary surgical and transcatheter interventions for establishment of right ventricular–pulmonary trunk continuity.

![Figure 6](http://heartjnl.com)
Concerns have been raised with regard to volume loading of the left ventricle and reduction of systemic diastolic pressure after shunt insertion. On the other hand, Hanley and colleagues reported that concomitant shunt insertion during RVOTR improves survival, possibly by preventing significant hypoxia during the early postoperative period.

Data in the literature that compare the outcomes of patients after surgical decompression of the RV with outcomes after transcatheter valvotomy are limited. In a recent study comparing the results of radiofrequency assisted valvotomy and balloon dilatation with CPV and Blalock-Taussig shunt, Alwi and colleagues concluded that transcatheter intervention is more effective in decompressing the RV and is associated with a lower morbidity. It is important to note, however, that patients selected for radiofrequency assisted valvotomy have relatively well developed tripapertite RVs. The present study, however, failed to confirm a lower reintervention rate after transcatheter opening of the RV outflow. This may be related to our inclusion of patients having moderately hypoplastic RVs (tricuspid valve z score as small as –3.42) for catheter intervention as reported previously.

The atretic pulmonary valve was successfully perforated by laser assistance in 94% (16 of 17) of our patients, which is comparable with the 75–100% reported by various investigators. By avoiding primary RV outflow surgery, catheter intervention may reduce the severity of RV damage and morbidity caused by postoperative complications. Nonetheless, perforation of the infundibulum and septis are not uncommon complications. At intermediate term follow up, we and others have shown that, although the absolute RV and tricuspid annulus dimensions increase after catheter intervention, RV area and tricuspid annulus z values decrease progressively with somatic growth, suggesting that a normal RV size may not be mandatory for sustaining a biventricular circulation. However, it remains unknown whether the RV growth is similar to that after RVOTR or CPV. Nonetheless, the proportion of patients who have achieved or are likely to achieve a biventricular circulation is similar between patients undergoing primary catheter intervention (11 of 14, 79%) and those undergoing primary RVOTR or CPV (34 of 43, 79%, p = 1.0). Our data compare favourably with the 64% (16 of 25) after catheter intervention as reported by Humpl and colleagues and the 76% (19 of 25) after surgery.

A number of limitations deserve comments. Firstly, the relatively small number of patients undergoing primary catheter intervention might have biased the outcome analysis. Even so, our findings concur with those of a recent study of 30 patients undergoing radiofrequency assisted valvotomy. Secondly, the management strategies have evolved over a period of two decades. Undoubtedly, the surgical mortality has been reduced significantly in the current era as illustrated by the results of Jahangiri and colleagues. However, even taking into account the improved radiofrequency assisted valvotomy with balloon valvoplasty proves to be an acceptable alternative. Further studies to compare the cost effectiveness and the severity of pulmonary regurgitation and its impact on long term RV function between surgical and catheter interventions may help to resolve the controversy.

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REFERENCES


IMAGES IN CARDIOLOGY

Double mycotic aneurysm of the descending thoracic aorta

A 71 year old man was admitted with fever and chills. On admission his temperature was 39°C; physical examination was normal except for a grade I/VI aortic systolic murmur. Blood cultures and urine samples grew Staphylococcus aureus. Transoesophageal echocardiography (TOE) showed a mildly calcified aortic valve without vegetations, and diffuse atheroma of the descending aorta. Despite appropriate antibiotic treatment the patient remained febrile, and developed clinical and radiological signs of pulmonary oedema. A subsequent TOE revealed the presence of two saccular aneurysms of the descending aorta, one just beneath the isthmus (panel A: AO, aorta), and the second one just above the diaphragm (panel B). These aneurysms arose from two localised ruptures in the atheromatous intima (arrows). The inferior one was responsible for a severe compression of the left atrium and right pulmonary veins, as shown by spectral and colour flow Doppler (panel C: LA, left atrium; LV, left ventricle; MA, mycotic aneurysm). Magnetic resonance imaging (spin echo T1 images) showed both mycotic aneurysms with a severely inflammatory shell and infiltration of the mediastinal fat, as well as the compressed left atrium (panel D: AA, ascending aorta; P.I.E, pleural effusion; RA, right atrium; RV, right ventricle; RPA, right pulmonary artery). The patient was considered too ill for surgery and died within a few days from multiorgan failure.

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