## RESEARCH

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# Impact of concomitant tricuspid annuloplasty on right ventricular remodeling in patients with rheumatic mitral valve disease



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## Abstract

**Background:** Studies on the management of functional tricuspid regurgitation (TR) during mitral valve operations have drawn inconsistent conclusions. This study was designed to compare the treatment strategy of concomitant tricuspid annuloplasty (TAP) against isolated mitral valve replacement (MVR) in rheumatic mitral valve disease patients, and to assess the effect of concomitant TAP on postoperative right ventricular (RV) remodeling and function.

**Methods:** One hundred-seventy patients with rheumatic mitral valve disease receiving MVR were categorized into TAP group (n = 124) and non-TAP group (n = 46). Clinical and echocardiographic data were collected preoperatively and at 1-year follow-up. Three-dimensional echocardiographic indices of RV geometry and function were analyzed.

**Results:** At baseline, concomitant TAP group had larger RV end-diastolic volume, more decreased RV ejection fraction and RV longitudinal strain than non-TAP group (all P < 0.001). At 1-year follow-up, TAP group had improved RV geometry and function. While adverse changes were observed in non-TAP group. In analysis of variance, the above indices demonstrated significant interaction with different treatment group (all P < 0.001). In multivariate regression analysis, independent of age and Maze procedure, concomitant TAP was associated with postoperative RV volume reduction (P < 0.001), improvement of RV ejection fraction (P < 0.001), and relieved postoperative functional TR severity (P = 0.025).

**Conclusions:** Our results suggest that concomitant TAP could improve RV remodeling and function for rheumatic mitral valve disease patients, while those with mild preoperative functional TR who had isolated MVR might experience RV dilation and deterioration of RV function at follow-up. Concomitant surgery for functional TR could be considered for patients undergoing MVR with rheumatic mitral valve disease.

**Keywords:** Right ventricular remodeling, Functional tricuspid regurgitation, Rheumatic mitral valve disease, Concomitant tricuspid valve annuloplasty

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## Introduction

Concomitant tricuspid annuloplasty (TAP) at the time of mitral surgery has gained growing acceptance recently. Despite surgical repair for severe tricuspid regurgitation (TR) at the time of left-sided valve surgery is a Class I recommendation under the current guidelines, the optimal management for moderate and mild functional TR depends more on whether the patient had dilated tricuspid annulus and/or right heart failure [1, 2]. Besides, patients with rheumatic heart disease had higher rates of developing late postoperative TR, those patients with preoperative significant TR demonstrate a survival benefit from concomitant TAP [3]. Rheumatic valve disease patients undergoing mitral valve replacement (MVR) might need more aggressive treatment for functional TR, but the threshold value for intervention is ambiguous. Previous studies mostly focused on degenerative and functional mitral valve disease, which have shown that for moderate and severe preoperative TR, concomitant TAP could improve postoperative right ventricular (RV) function, leading to favorable RV remodeling [4, 5]. However, data regarding the effect of TAP is inconclusive in patients with rheumatic mitral valve disease.

As RV is a crescent-shaped structure located in close proximity to the sternum, two-dimensional (2D) echocardiographic measurements of RV structure and function are often challenging. Especially for post-cardiac surgery patients, 2D-based assessment of longitudinal function may not be reflective of global RV function [6]. Three-dimensional (3D) echocardiography is well suited to evaluate RV volume and ejection fraction, and has been validated against cardiac magnetic resonance imaging [7]. Moreover, on the basis of full-volume data set, 3D speckle tracking echocardiography could provide comprehensive quantitative assessment of global RV myocardial function and avoid out-of-plane speckle motion in 2D imaging. Therefore, in this study, by the use of 3D echocardiography, we sought to evaluate the effect of TAP concomitant with MVR on RV remodeling and function for rheumatic mitral valve disease patients, and to compare the echocardiographic data of patients undergoing concomitant TAP with data of patients undergoing isolated MVR, preoperatively and at 1-year follow-up.

## Methods

#### Study population

This research protocol was compliant to the ethical guidelines of the 1975 Declaration of Helsinki, and was approved by the institutional review board of West China hospital. Between March 2016 to June 2017, 211 patients with rheumatic mitral valve disease scheduled for MVR provided their willingness to participate in this

study and signed informed consent. The diagnosis of rheumatic heart disease was according to the 2012 World Heart Federation criteria [8]. Patients had their clinical and echocardiographic assessment within 1 week before the operation. Thirteen patients were excluded for organic aortic valve or tricuspid valve (TV) disease, 10 for suboptimal echocardiographic window, 6 for coronary artery disease that needed revascularization, 1 for pacemaker implantation postoperatively. Five patients were lost to follow-up. At last, we enrolled 176 patients for the analysis.

All patients in this study underwent MVR, the use of mechanical mitral valve or bioprosthetic valve was according to patients' age and anticoagulant plan, and the addition of Maze procedure for patients with atrial fibrillation was according to each patient's informed preferences and the attending surgeon's discretion. All patients with moderate and severe TR had concomitant TAP. If TR grade was less than moderate, the implementation of TAP was at the attending surgeon's discretion. Tricuspid annular dilation (> 40 mm or > 21 mm/m<sup>2</sup>), atrial fibrillation, pulmonary hypertension and right heart failure were the main risk factors when considering whether less-than-moderate functional TR patients needed prophylactic TAP. However, some surgeons at our center would take rheumatic heart disease as a risk factor and perform prophylactic TAP for those with rheumatic mitral valve disease and less-than-moderate functional TR, irrespective of tricuspid annular dimension or pulmonary artery systolic pressure.

#### Echocardiographic and clinical data collection

Comprehensive transthoracic echocardiographic examinations were performed using the Philips EPIQ 7C (Philips Medical Systems, Andover, Massachusettes, USA) with a 2-MHz to 4-MHz matrix-array transthoracic transducer, which could display both 2D and 3D fullvolume images. Four consecutive cardiac cycles were recorded in patients with sinus rhythm and 6 cycles in atrial fibrillation. For patients with sinus rhythm, 3D full volume data set were obtained from 4 to 6 consecutive cardiac cycles during held respiration (frame rate  $\geq$  18 Hz). In patients with atrial fibrillation, high volume rate quantification mode was applied in the reconstruction of 3D full volume data. All the echocardiographic parameters were measure in 3 successive beats and then averaged.

LV volume and ejection fraction were calculated using Simpson's biplane method from 2D imaging. Left and right atrial volume were calculated by the disk summation technique from apical 4-chamber view. The tricuspid annulus diameter was measured from RV-modified apical 4-chamber view in end-diastolic and was indexed to body surface area. 2D RV systolic function was assessed by tricuspid annular plane systolic excursion (TAPSE), RV fraction area change and tissue Doppler of the tricuspid annulus. Systolic RV pressure gradient was calculated by the modified Bernoulli equation and then pulmonary artery systolic pressure (PASP) was derived. According to current guidelines [9, 10], TR severity was qualitatively graded as 4 categories: no or trace (jet area < 1 cm<sup>2</sup>), mild (vena contracta < 3 mm, jet area < 5 cm<sup>2</sup>), moderate (3 mm ≤ vena contracta < 7 mm), and severe (vena contracta ≥7 mm).

The 3D full volume images from RV-modified apical 4-chamber view were analyzed by the software of Tomtec Image Arena 4.0 on off-line Imaging workstation (TomTec Imaging Systems, Munich, Germany). After manual initialization of the RV endocardium at 8 points on the long axis and short axis views (Fig. 1, a), the software automatically traced the endocardium in the end-diastole and end-systole (Fig. 1, b). RV end-diastolic volume (RV EDV), RV end-systolic volume (RV ESV), RV ejection fraction (RVEF), RV longitudinal strain (RVLS) of free wall and septal wall were then calculated. The endocardial contours were manually adjusted to optimize the tracking of the myocardium. Negative strain values indicate myocardium contraction, reduced absolute values represented diminished contractile function. Reproducibility of 3D image analyses was determined in a random 10% of 3D images subgroup.

At baseline and during 1-year follow-up, symptoms were recorded and functional New York Heart Association (NYHA) class was assessed. Physical examination and electrocardiogram were performed before each echocardiographic assessment. Cardiopulmonary bypass time and other data of surgical procedure were extracted from medical records.

## Statistical analysis

Continuous variables were expressed as median and interquartile range, categorical variables were expressed as numbers and percentages. Comparison between 2 groups were performed by Mann-Whitney U test for



continuous variables, and  $\chi^2$  or Fisher's exact test for categorical variables as appropriate. Cumulative incidence rates of survival were estimated by the Kaplan-Meier method and compared using the log-rank test. To compare the effect of with and without concomitant TAP on cardiac remodeling between groups across the preoperative and 1-year follow-up periods, 2-factor analysis of variance with repeated measures were conducted. Within groups, changes of each continuous variable at follow-up time were analyzed by Wilcoxon signed rank test, and changes of dichotomous variables were analyzed by McNemar test. A sensitive analysis was performed in patients with less-than-moderate functional TR who received concomitant TAP, we compared their pre- and post-operative data with patient who had lessthan-moderate functional TR and isolated MVR.

To evaluate whether post-operative TR severity, RV remodeling and function were affected by multiple factors (including age, gender category, pre-operative PASP, concomitant Maze procedure, concomitant TAP), multivariate ordinal logistic regression was used to assess the risk factors for the severity of TR grade at 1-year follow-up, and multivariate linear regression was used to test the association between the abovementioned characteristics and percent change of postoperative RV EDV, RVEF, and RVLS.

Interobserver and intraobserver agreement for RV EDV, RVEF, RVLS of free wall and septal wall measurement were tested by Bland-Altman analysis (mean difference and standard deviation). Two independent blinded observers performed the offline analysis on 2 different days for 34 randomly selected images. Statistical analyses were conducted using the Statistical Package for Social Sciences release 25.0 (SPSS, Chicago, USA). All tests were 2-tailed, a P value < 0.05 was considered statistically significant.

## Results

Baseline characteristics and the effect of TAP at follow-up Among the 176 patients enrolled in this study, 6 patients were deceased within 6 months after the surgery, with 5 patients in the TAP group (3 patients deceased before discharge due to postoperative complication, 1 died of heart failure 1 month after the operation, 1 had sudden death 4 months after the operation, the cumulative survival rate was 96.1%), and 1 patient in the non-TAP group (due to stroke 6 months after the operation, the cumulative survival rate was 97.9%). No significant difference in all-cause mortality between the two groups was found (log-rank P = 0.57). Finally, 170 patients with complete 1-year postoperative clinical records and echocardiographic data were assessed (concomitant TAP group, n = 124; non-TAP group, n = 46). One hundred thirty-three patients had mechanical mitral valve and 37

had bioprosthetic valve. Concomitant tricuspid repair by DeVega annuloplasty was performed in 28 patients, and ring annuloplasty in 96 patients (Cosgrove-Edwards annuloplasty band with size of 26 mm, 28 mm, 30 mm and 32 mm). Table 1 demonstrated baseline characteristics and the operative information of the two groups. Compared with patients had isolated MVR, patients undergoing concomitant TAP were older, had smaller body size, worse cardiac function class, and higher prevalence of atrial fibrillation. With regard to echocardiographic data, preoperative TR grade were significantly greater in the TAP group, this group also had larger right heart volume, more dilated tricuspid annulus, and more decreased LV and RV function compared with non-TAP group at baseline (all P < 0.001). While patients in the isolated MVR group all had less-thanmoderate TR. Cardiopulmonary bypass time were longer in TAP group than the non-TAP group, and Maze procedure was more frequently performed in the TAP

group. Changes of clinical and echocardiographic data at 1year follow-up were summarized in Table 2, and the results of paired pre- and post-operative comparison as well as the interaction between groups and time were also showed. After surgery, the prevalence of atrial fibrillation was only significantly decreased in the TAP group. TR severity of the TAP group relieved at 1-year followup. However, we observed TR progression in the non-TAP group, in which 24% patients developed moderate and severe TR at follow-up. In terms of quantitative echocardiographic parameter, we considered the significant interaction between groups and time as the effect of concomitant TAP. The left heart volume and 2D parameters of RV did not show significant interaction, while estimated PASP and 3D RV parameters including RV EDV, RVEF and RVLS all showed significant interaction (all P < 0.001). Longitudinal changes of the 3D RV parameters were also demonstrated in Fig. 2. TAP group had worse baseline RV parameters, but during 1year follow-up, RV EDV reduced, RVEF and RV longitudinal strain improved. Whereas patients in non-TAP group had dilated RV, deteriorated RVEF and RVLS of free wall at follow-up. So that at 1 year after the surgery, there was no difference of RVEF or RVLS between the groups, while non-TAP group had slightly larger RV EDV (P = 0.01).

The results of sensitive analysis were presented in Fig. 3 and additional files. In patients with preoperatively less-than-moderate functional TR, 41% received prophylactic TAP concomitant with MVR (n = 32), the other 59% received isolated MVR (n = 46). Additional Table 1 demonstrated their baseline characteristics. TAP group was older, with more atrial fibrillation, and functional TR grade were more severe than the non-TAP group,

Variables	TAP group ( <i>n</i> = 124)	Non-TAP group $(n = 46)$	P value
Age, y	53.0 (47.0, 62.0)	50.0 (44.0, 59.0)	0.03
Female	98 (79%)	33 (72%)	0.32
Body surface area, m <sup>2</sup>	1.53 (1.45, 1.62)	1.59 (1.49, 1.66)	0.04
NYHA functional class III-IV	92 (74%)	22 (48%)	0.001
Atrial fibrillation	99 (80%)	16 (35%)	< 0.001
Echocardiographic data			
LV end-diastolic volume, ml	102.0 (86.3, 127.8)	100.5 (88.8, 120.0)	0.99
LV ejection fraction, %	61.0 (54.0, 66.0)	67.0 (60.0, 68.3)	< 0.001
Left atrial volume, ml	135.0 (99.0, 173.0)	119.5 (89.2, 138.0)	0.03
Right atrial volume, ml	51.1 (39.7, 67.0)	33.5 (25.8, 44.0)	< 0.001
TR grade			
None to trace	3 (3%)	17 (37%)	< 0.001
Mild	29 (23%)	29 (63%)	< 0.001
Moderate	57 (46%)	0	< 0.001
Severe	35 (28%)	0	< 0.001
TV annulus diameter, mm	34.0 (31.0, 36.8)	28.0 (27.0, 31.0)	< 0.001
Indexed TV annulus diameter, mm/m <sup>2</sup>	22.3 (20.6, 23.8)	18.2 (17.2, 19.6)	< 0.001
Pulmonary artery systolic pressure, mmHg	38.6 (33.0, 47.8)	29.8 (24.4, 38.0)	< 0.001
RV inlet dimension, mm	36.3 (32.9, 39.3)	32.9 (31.3, 37.0)	< 0.001
TAPSE, mm	11.9 (9.1, 15.2)	14.4 (10.4, 17.4)	0.01
RV fractional area change, %	40.2 (34.0, 44.9)	41.7 (36.0, 47.8)	0.09
Tissue Doppler of the tricuspid annulus, cm/s	9.2 (7.9, 10.7)	9.7 (8.2, 11.8)	0.15
RV end-diastolic volume, ml	51.3 (43.4, 64.6)	41.3 (35.8, 46.1)	< 0.001
RV ejection fraction, %	32.6 (28.6, 35.2)	40.9 (36.6, 46.1)	< 0.001
RVLS of free wall, %	-15.5 (-18.6, -13.4)	-22.0 (-24.3, -18.7)	< 0.001
RVLS of septal wall, %	-10.2 (-11.9, -8.6)	-12.3 (- 13.7, -10.9)	< 0.001
Operative data			
Concomitant MAZE procedures	59 (48%)	2 (4%)	< 0.001
Mechanical mitral prosthesis	94 (76%)	39 (85%)	0.21
Cardiopulmonary bypass time, min	100.0 (84.0, 126.8)	78.5 (66.8, 99.2)	< 0.001
MV lesion			
Stenosis	79 (64%)	34 (74%)	0.21
Regurgitation	17 (14%)	6 (13%)	0.91
Mixed	28 (23%)	6 (13%)	0.17

LV left ventricle, MV mitral valve, NYHA New York Heart Association, RV right ventricle, RVLS right ventricular longitudinal strain, TAP tricuspid annuloplasty, TAPSE tricuspid annular plane systolic excursion, TR tricuspid regurgitation, TV tricuspid valve

Data are expressed as median (interquartile range) or n (%)

also the preoperative RV function were worse in TAP group. But preoperative RV EDV was comparable between the two groups. Similar to the main analysis, at 1year follow-up, TR severity was worse in the non-TAP group. RV EDV, RVEF and RVLS of the septal wall demonstrated different changes in reverse direction according to group, TAP group had reduced RV EDV, increased RVEF, and improved RVLS of septal wall. On the contrary, non-TAP group showed dilated RV EDV and deteriorated RV function. For patients with lessthan-moderate TR, 3D parameters of RV EDV, RVEF, RVLS of free wall and septal wall showed significant interaction between surgery approaches and time. (Fig. 3 and additional Table 2).

	One year follow-up data		Compared with preoperative data		Two-way ANOVA	
	TAP ( <i>n</i> = 124)	Non –TAP ( <i>n</i> = 46)	P value	TAP ( <i>n</i> = 124)	Non –TAP ( <i>n</i> = 46)	 Interaction
NYHA functional class III-IV	21 (17%)	1 (2%)	0.009	< 0.001	< 0.001	-
Atrial fibrillation	54 (44%)	11 (24%)	0.02	< 0.001	0.13	-
Echocardiographic data						
TR grade						
None to trace	76 (61%)	11 (24%)	< 0.001	< 0.001	0.18	-
Mild	33 (27%)	24 (52%)	0.002	0.67	0.35	-
Moderate	14 (11%)	10 (22%)	0.09	< 0.001	0.002	-
Severe	1 (1%)	1 (2%)	0.47	< 0.001	1.00	-
LV end-diastolic volume, ml	102.0 (85.3, 112.0)	104.0 (90.5, 116.0)	0.42	0.06	0.54	0.76
LV ejection fraction, %	60.0 (58.0, 63.0)	61.0 (58.0, 62.3)	0.33	0.67	0.003	0.03
Left atrial volume, ml	86.9 (66.4, 114.0)	83.8 (57.8, 106.5)	0.15	< 0.001	< 0.001	0.63
Right atrial volume, ml	38.4 (31.7, 52.0)	38.6 (29.5, 44.6)	0.20	< 0.001	0.08	< 0.001
Pulmonary artery systolic pressure, mmHg	24.3 (20.7, 29.8)	24.7 (21.2, 28.1)	0.81	< 0.001	< 0.001	< 0.001
RV inlet dimension, mm	35.7 (32.9, 38.5)	35.8 (33.1, 37.3)	0.52	0.13	0.006	0.13
TAPSE, mm	12.3 (9.4, 14.3)	13.3 (11.2, 15.5)	0.03	0.42	0.06	0.27
RV fractional area change, %	42.3 (37.5, 46.6)	42.8 (38.4, 47.9)	0.35	0.005	0.16	0.44
Tissue Doppler of the tricuspid annulus, cm/s	8.6 (7.6, 9.4)	8.5 (7.9, 9.5)	0.77	< 0.001	< 0.001	0.24
RV end-diastolic volume, ml	44.5 (36.8, 51.4)	48.0 (42.9, 57.4)	0.01	< 0.001	< 0.001	< 0.001
RV ejection fraction, %	38.1 (34.5, 42.0)	38.6 (34.3, 42.4)	0.72	< 0.001	0.001	< 0.001
RVLS of free wall, %	-18.2 (-19.8, -15.9)	-19.2 (-20.7, -15.7)	0.32	< 0.001	< 0.001	< 0.001
RVLS of septal wall, %	-12.1 (- 15.8, -10.6)	-11.4 (-13.5, -8.9)	0.06	< 0.001	0.14	< 0.001

## Table 2 Clinical and echocardiographic data at 1-year follow-up

LV left ventricle, NYHA New York Heart Association, RV right ventricle, RVLS right ventricular longitudinal strain, TAP tricuspid annuloplasty, TR tricuspid regurgitation

Data are expressed as median (interquartile range) or n (%)

# Predictors of postoperative TR severity, RV remodeling and function

Table 3 showed the univariate and multivariate ordinal logistic regression analysis according to the grade of TR severity at 1-year follow-up. By univariate regression, increasing age was a risk factor for postoperative TR deterioration, while Maze procedure and TAP concomitant with MVR were protective factors for patients from TR worsening. Post-operative atrial fibrillation showed no associated with TR grade. In multivariate analysis, concomitant TAP remained as the independent predictor for the relief of postoperative TR (Odds Ratio: 0.45; 95% Confidence Interval: 0.22–0.91; P = 0.025). Predictors of longitudinal changes of RV volume and function by multivariate linear regression were demonstrated in Table 4. TAP was the main independent predictor of RV volume decrease at 1-year follow-up  $(R^2 = 0.41, \text{ Standardized } \beta = -0.557, P < 0.001). \text{ TAP}$ was also the independent predictor for RVEF increment  $(R^2 = 0.29)$ , Standardized  $\beta = 0.364$ , P < 0.001), as well as for improvement of RVLS of the free wall ( $R^2 = 0.28$ , Standardized  $\beta = 0.206$ , P = 0.01) and septal wall ( $R^2 =$ 0.14, Standardized  $\beta = 0.28$ , P = 0.001). In general, TAP was closely related to the RV reverse remodeling and functional improvement at 1-year follow-up. Meanwhile, concomitant Maze procedure was the independent predictor of the postoperative RVEF and RV strain improvement, but no significant correlation was found between Maze procedure and change of RV EDV at 1year follow-up.

#### Reproducibility

Inter-observer reproducibility by Bland-Altman analysis demonstrated good agreement: mean difference (standard deviation) was  $-0.06 (\pm 1.59)$  for RV end-diastolic volume,  $-0.21 (\pm 1.59)$  for RVEF, 0.35 ( $\pm 1.34$ ) and 0.16 ( $\pm 1.94$ ) for RVLS of the free and septal wall. Also good agreement was seen in intra-observer reproducibility:



0.01 ( $\pm$ 1.22) for RV end-diastolic volume, 0.18 ( $\pm$ 1.05) for RVEF, 0.24 ( $\pm$ 0.79) and 0.21 ( $\pm$ 0.85) for RVLS of the free and septal wall.

## Discussion

The major findings of the present study were the following: (1) Routine TAP at the time of MVR in rheumatic mitral valve patients could relieve residual functional TR, promote RV reverse remodeling and improve RV function in patients with worse baseline characteristics; (2) RV dilation and functional deterioration were observed in patients with isolated MVR at 1-year followup. These findings support the approach to perform TAP concomitant with MVR in rheumatic mitral valve disease patients. Prophylactic TAP could prevent RV dilation and functional deterioration postoperatively.

Patients with mitral valve disease often have functional TR, concomitant TAP with left heart operation is an effective treatment to reduce TR grade and improve RV function. Since increasing TR severity was reported to be independently associated with worse long-term survival, and functional TR could progress even after left heart valve surgery [11, 12], to repair severe TR concomitant with left heart surgery is now the routine practice. Also, patients with risk factors such as tricuspid

annular dilation and right heart failure could benefit from concomitant TAP even in the absence of severe TR. Moreover, previous studies reported that moderate TR per se was not benign, and concomitant tricuspid repair for those patients resulted in RV reverse remodeling, preventing postoperative TR progression [5, 13, 14]. Our findings resonated with the previous studies, in addition, 3D echocardiography and speckle tracking could provide a more detailed description of the RV. As TR severity is preload and afterload dependent and often inconstant, merely the TR severity or annular dimension might not be enough to identify patients at risk, especially when under anesthesia [15]. 2D parameters of RV function were less sensitive to detect post-operative changes compared with 3D measurements in our patients. Measurement of RV geometry and function by 3D speckle tracking could provide complementary information for follow-up evaluation, as well as for the decision-making process when considering TAP concomitant with mitral valve surgery. Besides, Ye et al. reported that RV morphology and function were more associated with postoperative prognosis than functional TR grade [16]. Kammerlander et al. also demonstrated that RV dysfunction is independently correlated with survival late after left heart valve procedure [17]. Our



observational study showed that TAP improved postoperative RV function, but no difference of mortality between TAP and non-TAP groups was found during 1year follow-up. Long-term follow-up is still in progress, and 3D echocardiography remains a convenient method for preoperative and follow-up assessment of RV morphology and function.

The postoperative deterioration of RV volume and function in patients with isolated MVR is a concern for surgeon to perform prophylactic TAP for less-thanmoderate functional TR. Sakata et al. had reported that in patients with moderate and mild TR who had no concomitant TAP, worsening of RV systolic function and dilation of RV diameter were prominent at 6-month to 1-year follow-up, while RV diastolic function was preserved [18, 19]. Similarly, Chikwe et al. demonstrated that RV dysfunction was prominent at discharge and around 1-year follow-up for patients undergoing mitral valve surgery [5]. Nevertheless, a random trial by Pettinari et al. revealed that, in patients with less-than-severe functional TR, prophylactic TAP irrespective of annular dilatation during mitral valve surgery could prevent significant postoperative TR, but had no effect on RV function and remodeling at 5-year follow-up [20]. The extrapolation of this randomized controlled trial should be cautious, as almost 60% of the patients had degenerative mitral disease, rheumatic valve disease patients might be under-represented in the trial. It had been observed that RV dilation and TR progression were rare after successful repair of degenerative mitral regurgitation without TAP [21]. However, in our patients of rheumatic mitral valve disease who had valve replacement rather than repair, postoperative RV dilation were seen in those with less-than-moderate preoperative TR receiving isolated MVR. We inferred that differences in the outcomes among our study and the previous studies probably due to the different etiologies of valvular disease. Rheumatic mitral valve patients are usually female, with higher prevalence of atrial fibrillation, having elevated PASP and lower body mass index compared with patients of degenerative mitral disease, which were risk factors of TR progression [22-24]. Even though MVR relieved LA overload and decreased pulmonary pressure, deterioration of TR grade and RV performance was still observed in our patients with mitral valve replacement. Besides the above risk factors, the possible cause might be attributed to the hypoxia during the operation, and prosthetic mitral valve might affect the shape and function of the tricuspid apparatus. Our result implied that prophylactic TAP could be considered for rheumatic

## Table 3 Predictors of post-operative TR severity at 1-year follow-up by ordinal logistic regression

	Univariate		Multivariate	
	P value	OR (95% CI)	P value	OR (95% CI)
Age	0.02	1.03 (1.01–1.07)	0.17	1.02 (0.99–1.05)
Female	0.89	1.05 (0.53–2.08)	-	_
Pre-operative PASP	0.88	0.99 (0.98–1.02)	-	_
Post-operative atrial fibrillation	0.14	1.56 (0.87–2.82)	-	-
Concomitant Maze procedure	< 0.001	0.21 (0.10-0.41)	0.001	0.29 (0.14–0.61)
Concomitant TAP	< 0.001	0.27 (0.14–0.52)	0.025	0.45 (0.22–0.91)

CI confidence interval, OR odds ratio, PASP Pulmonary artery systolic pressure, TAP tricuspid annuloplasty, TR tricuspid regurgitation

mitral valve disease patients with less-than-moderate functional TR.

Whether adding TAP to left heart surgery for mild or trace TR could improve postoperative survival remains to be clarified. Kim et al. found that, in rheumatic mitral valve patients, risk of death and congestive heart failure were similar between TAP and non-TAP group for mild-to-moderate TR [25]. In another study of 959 patients with mild-to-moderate TR, concomitant TAP was not associated with better clinical outcomes, but resulted in reduced postoperative TR grade [26]. In the research by Lee et al. of patients with less-than-moderate TR undergoing MVR, TAP showed a tendency to prevent TV-related events in a median follow-up duration of 8 years, notably 93% of their patients had rheumatic etiology [27]. Besides, positive effect of prophylactic TAP on valve-related events was found in rheumatic valve disease patients with less-than-moderate TR undergoing double valve replacement [28]. Whether patients with more rheumatic burden could benefit more from

**Table 4** Multivariate linear regression analysis of the predictors for post-operative right ventricular remodeling and function in rheumatic valve patients undergoing mitral valve replacement

Dependent variables	Independent variables	Standardized $\beta$	P value
Percent change of RV EDV at 1-year follow-up	Age	0.004	0.95
	Female	0.045	0.45
	Pre-operative PASP	0.163	0.01
	Concomitant Maze procedure	-0.027	0.69
	Concomitant TAP	-0.557	< 0.001
Percent change of RVEF at 1-year follow-up	Age	0.102	0.14
	Female	0.004	0.95
	Pre-operative PASP	0.026	0.71
	Concomitant Maze procedure	0.212	0.004
	Concomitant TAP	0.364	< 0.001
Percent change of free wall RVLS at 1-year follow-up	Age	0.149	0.03
	Female	-0.051	0.45
	Pre-operative PASP	0.059	0.41
	Concomitant Maze procedure	0.342	< 0.001
	Concomitant TAP	0.206	0.01
Percent change of septal wall RVLS at 1-year follow-up	Age	0.01	0.90
	Female	-0.02	0.74
	Pre-operative PASP	0.10	0.20
	Concomitant Maze procedure	0.11	0.18
	Concomitant TAP	0.28	0.001

EDV end diastolic volume, PASP Pulmonary artery systolic pressure, RV right ventricle, RVEF right ventricular ejection fraction, RVLS right ventricular longitudinal strain, TAP tricuspid annuloplasty, TR tricuspid regurgitation

prophylactic TAP concomitant with left heart surgery was a debated topic. Although concomitant TAP had positive effect on RV geometry and function in our rheumatic patients, the long-term clinical effect still need to be assessed. Even with less-than-moderate TR, rheumatic valvular disease patients receiving concomitant TAP were older, with atrial fibrillation, and decreased RV function compared with the isolated MVR group. Nevertheless, concomitant TAP improved RV geometry and function at 1-year follow-up in our patients with worse baseline risk factor. Considering the safety of performing additional TAP concomitant with left heart surgery and the risk of reoperation for severe TR [29], we incline to suggest prophylactic TAP concomitant with MVR for rheumatic mitral valve disease to prevent RV dilation and TR progression at follow-up.

There were several limitations in our study. First, as its observational nature, patients were treated by several surgeons in whom different standards for operation and different surgical technique existed, some patients with less-than-moderate TR in our study had no guidelinerecommended indication of concomitant TAP, for whom prophylactic TAP was based on the attending surgeon's discretion. This selection bias and the potential confounder brought by different surgeons was inevitable. Second, the power of tests could be affected as the sample size was limited. Due to the limited sample size of patients with less-than-moderate TR, propensity score matching could not achieve sufficient pairs of patients from concomitant TAP and non-TAP group, and the comparison was underpowered. The effect of concomitant TAP in rheumatic valvular disease patients with less-than-moderate TR need to be tested by randomized control trial or large observational cohort. Third, there was no analysis of the RV geometry and function immediately postoperative, thus the trends of RV remodeling after the surgery were incomplete. Another limitation was that, due to the short-term follow-up of 1 year, we were not able to detect the long-term clinical outcome of concomitant TAP, or whether postoperative RV reverse remodeling could reduce valve-related adverse cardiac events. Further studies with large sample size and long-term follow-up is required.

## Conclusion

In this study focused exclusively on rheumatic mitral valve disease patients, we found that concomitant TAP with MVR had positive effect on postoperative RV geometry and function, independent of concomitant Maze procedure. Whereas patients with isolated MVR had RV dilation and deterioration of RV function at 1-year follow-up. Our results support that rheumatic mitral valve patients undergoing MVR could benefit from concomitant surgery for functional TR.

#### Abbreviations

2D: Two-dimensional; 3D: Three-dimensional; EDV: End-diastolic volume; EF: Ejection fraction; ESV: End-systolic volume; LV: Left ventricle; MV: Mitral valve; MVR: Mitral valve replacement; NYHA: New York Heart Association; RV: Right ventricle; RVLS: Right ventricular longitudinal strain; TAP: Tricuspid annuloplasty; TR: Tricuspid regurgitation; TV: Tricuspid valve

#### **Supplementary Information**

The online version contains supplementary material available at https://doi. org/10.1186/s12947-021-00245-2.

Additional file 1: Additional Table 1. Preoperative characteristics of patients with less-than-moderate functional tricuspid regurgitation. Additional Table 2. Follow-up data in patients with less-than-moderate functional tricuspid regurgitation

#### Acknowledgements

None

#### Authors' contributions

Li Rao conceived the study design, performed the final approval of the version to be submitted. Yue Zhong made the statistical analysis and drafted the manuscript. Wenjuan Bai checked the analysis results and revised the manuscript carefully. Hui Wang and Hong Qian participated in the enrollment of patients, and analyzed the echocardiographic image. All authors read and approved the final manuscript.

#### Funding

This study was supported by Science & Technology Pillar Program of Sichuan Province (No. 2017FZ0077), Post-Doctor Research Project, West China Hospital, Sichuan University (No. 19HXBH042) and 1-3-5 project for disciplines of excellence-Clinical Research Incubation Project, West China Hospital, Sichuan University (No. 2020HXFH044).

#### Availability of data and materials

All data generated and analyzed in this study is available from the corresponding author on reasonable request.

#### Declarations

#### Ethics approval and consent to participate

This study was authorized by the ethics committee of West China Hospital of Sichuan University, and written informed consent was obtained from all the participates.

#### Consent for publication

Consent for publication was obtained from all the participates.

#### **Competing interests**

All authors declare that they have no competing interests.

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#### Received: 27 December 2020 Accepted: 25 February 2021 Published online: 04 March 2021

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