The role of strut chordae in mitral valve competence during annular dilation

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Abstract
Strut chordae, on their own, are not typically thought to aid mitral valve competence. The aim of this study is to assess whether strut chordae aid mitral valve competence during acute annular dilation. Twelve porcine hearts were dissected and tested using an in vitro simulator, with the mitral annulus tested in either a ‘normal’ or a dilated configuration. The normal configuration included a diameter of 30 mm, a posterior leaflet ‘radius’ of 15 mm and a commissural corner ‘radius’ of 7.5 mm; the dilated annular template instead used dimensions of 50 mm, 25 mm and 12.5 mm, respectively. Each mitral valve underwent ten repeat tests with a target systolic pressure of 100 mmHg. No significant difference in the pressure was detected between the dilated and regular annuli for the mitral valves tested (95 ± 3 mmHg cf. 95 ± 2 mmHg). However, the volume of regurgitation for a dilated annulus was 28 ml greater than for a valve with a normal annulus. Following severing of strut chordae, there was a significant reduction in the systolic pressure withstood before regurgitation by mitral valves with dilated annuli (60 ± 29 mmHg cf. 95 ± 2 mmHg for normal annular dimensions; p < 0.05). In conclusion, strut chordae tendineae may play a role in aiding mitral valve competence during pathophysiology.

Keywords
annular dilation; chordae tendineae; mitral valve; regurgitation; strut chordae

Introduction
The mitral valve is composed of the anterior and posterior leaflets and chordae tendineae which connect these leaflets to papillary muscles and in turn the left ventricle.1 During systole, the leaflets have to cover an orifice, the circumference of which is outlined by a ring of tissue known as the mitral annulus. Closure of the mitral valve leaflets during systole prevents mitral regurgitation, enabling blood to flow out of the aortic valve and through to the aorta. At this stage, valve competence is maintained by coaptation of the two leaflets; that is, they close by pressing against each other and so prevent blood from flowing back through the mitral valve. During diastole, the leaflets open to enable blood to flow into the left ventricle from the left atrium. The anatomy and biomechanics of the mitral valve, and its subvalvular apparatus, are reviewed elsewhere in more detail.2

Mitral valve regurgitation is a condition prevalent in up to 21.3% of the population.3 Although severe mitral regurgitation is less prevalent, it can require surgical correction to reduce the risk of morbidity and mortality. A known cause of mitral valve regurgitation is mitral valve annular dilation,2,4 which can be associated with dilated cardiomyopathy.5 The result is impaired left ventricular outflow of blood during systole due to mitral insufficiency.

Mitrval regurgitation can be caused by a failure of chordae tendineae of the mitral valve. Categories of chordae tendineae include marginal and basal chordae. Strut chordae are a type of basal chord and are the largest and thickest chordae of the mitral valve. They insert into the anterior leaflet, linking the anterior leaflet to papillary muscles (Figure 1). Strut chordae are currently believed to have very little effect on regurgitation when ruptured;5–8 however, rupture of marginal chordae is known to lead to mitral regurgitation.5,6,9 This is despite strut chordae carrying a greater load than marginal chordae.10 Instead, strut chordae are thought to enable

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physiological motion of the anterior leaflet.\textsuperscript{7,8,11} However, this may relate to the greater extensibility of strut chordae than marginal chordae\textsuperscript{12,13} which when combined with the differing load transfer may enable smooth closure of the mitral valve. It is unknown, though, whether during annular dilation, strut chordae have a greater role in valve closure.

The aim of this study is to assess whether strut chordae aid mitral valve competence during acute annular dilation. An \textit{in vitro} simulator\textsuperscript{16–18} has been used so that valve competence could be analysed in both regular and dilated annuli, with strut chordae intact and severed. The advantage of using an \textit{in vitro} simulator is that it enables precise test conditions to be mimicked. In this study, porcine hearts have been used, as they are anatomically similar to humans\textsuperscript{19} with regard to their size, shape of leaflets and arrangement of chordae tendineae.\textsuperscript{20}

\section*{Methodology}

\subsection*{Specimens}

Twelve porcine hearts were obtained from a supplier (Samples for Schools, Littlehampton, UK). Each heart was individually wrapped in paper towels, soaked in Ringer’s solution, subsequently placed in heat sealed bags and frozen at $-40^\circ$C until required for dissection.\textsuperscript{6,16} Prior to dissection, each heart was kept at $4^\circ$C for 24 hours to ensure it was fully thawed. The dissection procedure is further detailed elsewhere,\textsuperscript{16} but briefly, the right side of the heart was removed along with the apex of the left ventricle. An incision was made into the ventricle cavity to expose the subvalvular apparatus including chordae tendineae (Figure 2).

Following dissection, the hearts underwent a second freeze cycle, until required for testing. Ahead of testing, dissected mitral valves were sutured onto a rubber sheet to mimic either the dimensions of a ‘normal’ mitral annulus or a dilated annulus (Figure 3). Two anchor stitches were used to align the ‘D’-shaped hole (Figure 3), with the annulus then sutured along the circumference of the rubber template (Figure 3). The normal mitral annulus template had an inter-commissural diameter of 30 mm, a posterior leaflet ‘radius’ of 15 mm and a commissural corner ‘radius’ of 7.5 mm (Figure 3) based on suggested ratios from literature.\textsuperscript{21} For the dilated template, these dimensions were increased to 50 mm, 25 mm and 12.5 mm, respectively.

\subsection*{In vitro mechanical testing}

The \textit{in vitro} simulator used to test heart valves is described extensively elsewhere.\textsuperscript{16} Briefly, the simulator was connected via flexible tubing onto a pump used to
pressurise water (Model 12,107-15, Cole-Parmer, London) (Figure 4). Pressure was monitored using a pressure transducer (Omega Engineering Limited, Northbank Ind Park, Manchester, UK), noting that 1 mmHg = 0.1333 kPa. The *in vitro* simulator consists of an outer cylinder, an adjustable inner cylinder and a clamping device, which holds a papillary muscle ‘annulus’ within an adjustable inner cylinder. The clamping device consists of a small section of cylinder mounted to a disc. This cylinder is inserted into the ventricle via the exposed apex. Two plates of Perspex, one on the posterior and the other on the anterior side of the leaflet, hold down the cylinder maintaining the papillary muscles in place.

The pressure and volume of regurgitation was measured for mitral valves. Six mitral valves were tested with no annular dilation (i.e. using the ‘normal’ annulus template) and a separate six mitral valves tested with annular dilation (i.e. using the dilated annulus template). Each mitral valve underwent ten repeat tests with target pressures of 100 mmHg to match the mean porcine arterial pressure of 102 mmHg. The first test ran for approximately 15 s, leading to repeatable pressure measurements, following which 9 repeat tests were performed. Following testing of each intact mitral valve (either with no dilation or with dilation), strut chordae were severed, with the testing procedure repeated. However, following the severing of chordae, the initial test had to run for approximately 35 s, in addition, as these valves leaked severely before initial coaptation (following severing of strut chordae), the volume of mitral regurgitation was not measured. A sample set of pressure–time data is provided in Figure 5 (following the severing of chordae).

**Data analysis**

Both a peak pressure and a stabilised pressure were obtained for each of 240 individual tests performed. Results from the first three tests were excluded when...
calculating mean and median pressures to negate any potential effects of pre-conditioning on the measurements made.

All statistical analyses were performed using Minitab (Version 18, Minitab Ltd, Progress Way, Coventry, UK). A normality test was used to determine whether data were normally distributed (Kolmogorov–Smirnov). While the peak and stabilised pressures of the dilated annulus were normally distributed, along with the peak pressures for tests on the regular annulus (p > 0.05), the stabilised pressures for the regular annulus were not normally distributed (p < 0.05). Therefore, comparisons for stabilised pressures of regular annuli used a non-parametric test, namely, a Wilcoxon signed rank test for chordal cutting and a Mann–Whitney test for annular dilation. All other statistical comparisons were performed using parametric tests (i.e. a paired T-test for chordal cutting and two sample T-tests for annular dilation). All significant differences were assessed to a 5% level (p < 0.05).

Results

Figure 6 outlines the key data obtained during this study. There was no significant difference in the stabilised pressure between the dilated and regular annuli for the mitral valves tested (95 ± 3 mmHg cf. 95 ± 2 mmHg, respectively; p > 0.05, Table 1). The peak pressures reached for the regular annulus (109 ± 7 mmHg) were not significantly higher than the peak pressures for valves with dilated annuli (105 ± 6 mmHg; p > 0.05, Table 2). However, the volume of regurgitation for a dilated annulus was 28 ml greater than for a valve with a normal annulus, a significant increase (p < 0.05; Table 3).

There was no significant difference in stabilisation (p > 0.05) or peak (p > 0.05) pressure for mitral valves with normal annuli before (95 ± 3 mmHg and 109 ± 7 mmHg, respectively) or after (94 ± 8 mmHg and 104 ± 12 mmHg, respectively) severing strut chordae (Table 1 and Table 2). However, when the strut chordae
were severed, there was a significant reduction in both stabilised (95 ± 2 mmHg vs 60 ± 29 mmHg, respectively; p < 0.05, Table 1) and peak pressure (105 ± 6 mmHg vs 63 ± 31 mmHg, respectively; p < 0.05, Table 2) for the dilated annuli.

### Table 1. Descriptive statistics for stabilised pressures in both the dilated and regular annuli before and after severing strut chordae (*significant difference before/after severing chordae; p < 0.05).

| Annulus | Intact (mmHg) | Severed chordae (mmHg) |
|---------|---------------|------------------------|
|         | Mean  | SD    | Median | Mean  | SD   | Median |
| Regular |       |       |        |       |       |        |
| 1       | 96.58 | 1.61  | 96.77  | 97.79 | 0.80  | 98.02  |
| 2       | 95.86 | 1.18  | 95.85  | 95.86 | 1.09  | 95.99  |
| 3       | 96.52 | 0.49  | 96.60  | 94.80 | 2.24  | 95.44  |
| 4       | 96.72 | 0.66  | 96.80  | 98.83 | 1.08  | 98.33  |
| 5       | 92.11 | 0.96  | 92.43  | 97.64 | 0.90  | 97.68  |
| 6       | 89.33 | 0.72  | 89.40  | 77.32 | 1.34  | 76.66  |
| Mean    | 94.52 | 3.09  | 96.22  | 93.71 | 8.16  | 96.83  |

| Dilated |       |       |        |       |       |        |
|---------|-------|-------|--------|-------|-------|--------|
| 1       | 95.53 | 0.61  | 95.72  | 88.31 | 1.20  | 88.18  |
| 2       | 94.89 | 0.56  | 95.09  | 41.14 | 0.55  | 41.04  |
| 3       | 95.64 | 0.95  | 95.58  | 81.62 | 1.36  | 81.34  |
| 4       | 93.27 | 1.56  | 92.72  | 83.06 | 0.91  | 83.16  |
| 5       | 97.13 | 0.43  | 97.09  | 17.14 | 1.16  | 17.75  |
| 6       | 92.99 | 0.48  | 92.91  | 48.10 | 2.76  | 47.38  |
| Mean    | 94.91*| 1.56  | 95.33  | 59.89 | 28.76 | 64.36  |

**SD:** standard deviation.

### Table 2. Descriptive statistics for peak pressures in both the dilated and regular annuli before and after severing strut chordae (*significant difference before/after severing chordae; p < 0.05).

| Annulus | Intact (mmHg) | Severed Chordae (mmHg) |
|---------|---------------|------------------------|
|         | Mean  | SD    | Median | Mean  | SD   | Median |
| Regular |       |       |        |       |       |        |
| 1       | 116.68 | 1.36  | 117.22 | 105.42 | 0.98  | 105.28 |
| 2       | 111.58 | 1.52  | 111.51 | 108.58 | 1.22  | 108.80 |
| 3       | 114.06 | 1.56  | 114.83 | 113.22 | 2.10  | 114.04 |
| 4       | 104.14 | 1.96  | 104.18 | 105.71 | 1.13  | 105.63 |
| 5       | 108.77 | 2.72  | 108.90 | 109.66 | 1.76  | 108.59 |
| 6       | 97.91 | 1.82  | 98.79  | 79.69 | 1.84  | 79.29  |
| Mean    | 108.86 | 6.89  | 110.20 | 103.71 | 12.11 | 107.11 |

| Dilated |       |       |        |       |       |        |
|---------|-------|-------|--------|-------|-------|--------|
| 1       | 107.78 | 1.52  | 107.54 | 97.64 | 1.98  | 96.89  |
| 2       | 98.65 | 0.58  | 98.94  | 42.67 | 0.54  | 42.47  |
| 3       | 104.67 | 2.63  | 104.75 | 85.46 | 1.43  | 85.13  |
| 4       | 101.07 | 1.98  | 100.43 | 86.76 | 1.61  | 86.88  |
| 5       | 115.32 | 0.97  | 115.35 | 18.43 | 1.28  | 19.06  |
| 6       | 103.20 | 3.34  | 105.06 | 49.14 | 3.20  | 48.27  |
| Mean    | 105.11*| 5.89  | 104.90 | 63.35 | 31.18 | 66.70  |

**SD:** standard deviation.

### Table 3. The mean, median and standard deviation of the pre-valve coapting regurgitation for both annulus sizes when the chordae are uncut (*significant difference between regular and dilated annulus; p < 0.05).

| Annulus | Mean | SD  | Median |
|---------|------|-----|--------|
| Regular |      |     |        |
| 1       | 6.71 | 3.73| 6      |
| 2       | 27.86| 8.53| 25     |
| 3       | 6.00 | 2.31| 5      |
| 4       | 6.71 | 1.38| 6      |
| 5       | 6.57 | 1.72| 6      |
| 6       | 23.14| 4.45| 24     |
| Mean    | 12.83| 9.93| 6      |

| Dilated |      |     |        |
|---------|------|-----|--------|
| 1       | 31.29| 7.14| 34     |
| 2       | 15.43| 2.57| 15     |
| 3       | 38.86| 8.53| 38     |
| 4       | 53.29| 8.54| 55     |
| 5       | 48.57| 4.86| 49     |
| 6       | 56.29| 11.56| 53    |
| Mean    | 40.62*| 15.45| 43.5 |

**SD:** standard deviation.

### Discussion

This study has provided initial evidence that strut chordae may have a role in maintaining some level of mitral valve competence during mitral annular dilation. Using an *in vitro* simulator, a peak pressure, stabilised pressure and regurgitation volume have been measured to allow for a comparison of systolic performance. When the strut chordae were severed in valves without mitral annular dilation, there was no change in performance, which is consistent with previous studies.6,16,23 Annular dilation did increase the regurgitation volume during initial coaptation, which is again consistent with results from literature.4,24 Although annular dilation alone did not significantly reduce either a peak or stabilised pressure, in combination with severed strut chordae it resulted in a significant drop in both peak and stabilised pressures. Indeed, mean stabilised pressures and peak pressures for ‘normal’ valves (95 mmHg and 109 mmHg, respectively) were not altered after strut chordae were severed (94 mmHg and 104 mmHg, respectively); these values are also within range of a target porcine physiological pressure of 102 mmHg.22

Valves with annular dilation experienced a significant loss in both stabilised and peak pressure when strut chordae were severed. It should be noted that many studies in literature which assess severing of strut chordae focus on models of ischaemic mitral regurgitation4,25,26 including with annular dilation.27 Comparisons across such studies are limited because fundamentally they are...
different studies. Studies on ischaemic mitral regurgitation include tethering of mitral leaflets due, potentially, to an imbalance in tension within chordae. Therefore, severing a strut chord would restore some form of functional valve competence. In this study, such tethering was not mimicked; instead, valve competence has been compromised by annular dilation, with a significant effect on valve competence noticeable following the severing of strut chordae.

Clinically, there is evidence that preserving the subvalvular apparatus improves results following mitral valve replacement surgery. The results from this study are supportive of such findings. Thus, with the advent of trans-catheter heart valve replacement, including its potential application to the mitral valve, our study would support the deployment of the valve while maintaining the native subvalvular mitral apparatus. The key finding from this study is that strut chordae may aid not just the mechanism of closure, but also valve competence during pathophysiology. Strut chordae tendineae may play a role in aiding mitral valve competence during pathophysiology. Strut chordae may aid marginal chordae in the prevention of mitral regurgitation in valves which are afflicted by mitral annular dilation.

Conclusions

Strut chordae tendineae may play a role in aiding mitral valve competence during pathophysiology. Strut chordae may aid marginal chordae in the prevention of mitral regurgitation in valves which are afflicted by mitral annular dilation.

Statement involving human and animal rights

This study did not involve any human participants or animal studies, and no animals were sacrificed specifically for this study. It is noted that animals from which porcine hearts were obtained were otherwise destined for the food chain. Ethical approval was granted for this study by the University of Birmingham Research Support Group [ERN_15-0,032].

Declaration of Conflicting Interests

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