

# Measuring chordae tension during transapical neochordae implantation: Toward understanding objective consequences of mitral valve repair

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1	Measuring chordae tension during transapical neochordae implantation:
2	towards understanding objective consequences of mitral valve repair
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32 **Conflict of interest** for this work: no conflict for any authors. 33 34 **Disclosure**: 35 Daniel GRINBERG: N/A 36 Pierre-Jean COTTINET: N/A 37 Florent GANET: N/A 38 Matteo POZZI: N/A 39 Sophie THIVOLET: N/A 40 David AUDIGIER: N/A 41 Jean-Fabien CAPSAL: N/A 42 Minh-Quyen : N/A 43 Jean-François OBADIA : 44 Resarch support: Boeringher, Saint Jude Medical, Abbott, Medtronic, Edwards -Consulting Fees/Honoraria: Edwards, Saint Jude Medical, Medtronic, Servier, Novartis -45 \_ \_ Royalty Income: Landanger, Delacroix-Chevalier 46 47 48 **Funding**: private funding 49 50 Corresponding author's complete contact information 51 Daniel GRINBERG, 52 Sinai BioDesign 53 Icahn School of Medicine at Mount Sinai Hospital 54 55 1428 Madison Ave, BMC-24 New York NY 10029 56 57 USA 58 59 Phone: +1 (646)-683-8599 60 Fax : +1 (212)-659-6818 Email: daniel.grinberg.pro@gmail.com - daniel.grinberg@mountsinai.org 61 62 63 Clinical trial registry number: N/A (Ethical committee certificate linked) 64 65 Article word count : 3393 / 3500 66 67 **Glossary Abbreviations** 68 69 MVr = Mitral valve repair 70 DMR = degenerative mitral regurgitation 71 MR = mitral regurgitation 72 TEE = transesophageal echocardiography73 3D-TEE = 3-dimentional TEE

# 74 <u>Central Picture (83 characters)</u>

75 Real-time choral tension measurements during a transapical neochordae implantation



# 77 <u>Central Message (199 characters)</u>

- 78 Tension applied on chordae sustaining P2 segment was 0,8 Newton and was spread homogeneously to
- all chordae. When normal physiology is restored, the subvalvular apparatus was in a low-stress state.
- 80

# 81 <u>Perspective (382 characters)</u>

- 82 We experienced a surgical strategy based on objective, quantitative parameters instead of only
- 83 echocardiogram.
- 84 New tools are needed to explore hypotheses raised by this work.
- 85 The prognostic value of the measured tension has to be evaluated.
- 86 Our protocol is more precise and practical than the current standard for final adjustment of chordae
- 87 length and could be used even without tension assessment.

#### 88 Structures Abstract (250 words)

89 <u>Objectives:</u>

90 Complex structure of mitral valve and its central position in the heart limit assessment of mitral 91 function to standardized calculated parameters assessed by medical imaging (echocardiography).

Novel techniques, which allow mitral repair (MVr) on a beating heart, offer the opportunity for
innovative objective assessment in physiologic and pathologic conditions. Here we report the first

94 data of real-time choral tension measurement during a transapical neochordae implantation.

95 <u>Methods:</u>

Seven patients with severe degenerative mitral regurgitation (DMR) due to posterior prolapse
underwent transapical MVr using NeoChord®. During prolapse correction, the tension applied on the

98 neochordae was measured; in addition to hemodynamic and echocardiographic parameters.

99 <u>Results</u>

100 The traction applied on one chorda sustaining the P2 segment was measured between 0.7 and 0.9101 Newtons, and oscillated with respiration.

102 When several neochordae were set in tension, this initial tension was spread homogeneously on each 103 chorda (mean sum of the amplitude of tension  $0.98 \pm 0.08$  N).

To achieve an optimal echocardiographic correction, a complementary synchronous traction on all chordae was required. During this adjustment, the sum of the tension decreased (mean  $12 \pm 2$  %, p = ,018), suggesting that when normal physiology is restored, the valvular apparatus was in a low-stress state.

108 This method allowed us to apply a precise and reproducible technique, leading to a good procedural 109 success rate with a low morbidity-mortality rate.

110 <u>Conclusions</u>

111 The tension applied on chordae during transapical implantation of neochordae for DMR can be

112 measured, providing original data about the objective consequences of MVr on the mitral apparatus.

# 113 I. Introduction

Mitral valve repair (MVr) remains the gold standard for treating primary mitral regurgitation (MR)<sup>1</sup>.
The technique has since evolved, and today the trend is to use more Gore-Tex to replace the elongated or ruptured chordae and to resect less tissue, thereby creating a larger surface of coaptation<sup>2</sup>. One of the main difficulties of this technique is to define the appropriate length of the chordae assessed on an empty non-beating heart.
A change of paradigm recently occurred with the introduction of novel techniques performed on

120 beating hearts like transapical neochordae using the Neochord DS 1000 (NeoChord Inc., Minneapolis,

121 Minnesota, USA). These devices allow real-time adaptation of the repair thanks to an intraoperative

122 transesophageal echocardiography (TEE) control.

We designed a device that enables a precise traction control on each chorda to determine the optimal length of the chordae during the NeoChord implantation, as well as measuring the tension applied on neochordae along this traction.

126 We also recorded original results on chordae tension *in vivo* that have not been previously described to

127 our knowledge, illustrating the physiology and the pathophysiology of the mitral valve.

#### 129 II. Methods

#### 130 II.1 PATIENTS

Since the use of NeoChord is not yet included in international guidelines, it was only proposed to patients with severe symptomatic MR due to a posterior prolapses, involving mainly P2 segment (with a variable extension on P1 or P3, but never involving the commissures or the anterior leaflet) and considered as high-risk for conventional surgery (mainly due to frailty) after selection by our heart team.

136 Informed consent was obtained from each patient about the complete procedure and to acknowledge 137 that their medical records will be collected in a registry.

This procedure has received the approval of the Ethical Committee of the French Society ofCardiology.

#### 140 II.2 ORIGINAL OPERATIVE TECHNIQUE

The usual NeoChord implantation protocol<sup>3</sup> can be summarized as: access to the left ventricular apex; establishment of two apical purse strings for the introduction of the device after systemic heparinization; implantation of several neochordae through the free edge of the prolapsed leaflet under 3-dimentional TEE (3D-TEE) control; fine adjustment of neochordae length by manual traction on each chordae to achieve the best coaptation under TEE control; fixing the chordae at the apex of the left ventricle at the predefined optimal length; protamine infusion, drainage, and closing.

147 Our conventional institutional anesthesiologic protocol was followed in order to maintain a mean148 blood pressure between 70 and 80 mmHg.

## 149 II.3 <u>MODIFICATION OF THE TECHNIQUE</u>

150 Using the original technique, the adjustment of the tension is delicate and approximate since it is 151 performed manually.

152 To improve the safety, the precision, and the reproducibility of this adjustment, we developed and

153 patented a specific adjustment device in collaboration with a team of engineers (LGEF at INSA-Lyon) 154 (Figure 1). The device consists of four identical units. Each unit consists of a crocodile clip system 155 connected to a force sensor (the force range and the resolution of the load cell are  $\pm$  10 N and 0.01 N 156 respectively) held on a manual linear axis system (micrometric leadscrew). It is, therefore, possible to 157 independently adjust the transverse tension exerted on each chorda connected to the force device. The 158 four units are fixed on a 120-mm-stroke-length linear table. Such a configuration enables the system 159 to apply the same mechanical traction to all four chordae synchronously using the principal screw. 160 The whole device remains lightweight, compact, and easy to transport.

161 Three to four NeoChords are usually required to hold the entire free-edge of the prolapsed segment, 162 and this statement can by validated by performing a brief manual traction on all chordae under TEE 163 control.

164 After this step the measurement tool is brought closer to the patient. The implanted neochordae are 165 then connected to the force device through the crocodile clips.

The measurement procedure consisted of four steps, as follows (Table 1): (1) The chorda implanted in the best location (in the middle of the prolapse) was tracked alone until the best TEE control; (2) The other chordae were then tracked, one at a time, with an individual screw to achieve an equivalent tension on all chordae; (3) All neochordae were then tracked together under TEE control, until obtaining the optimal MVr (no residual MR, vanishing of leaflet billowing, and homogeneous and high coaptation height); (4) the chordae are then fixed at the apex of the left ventricle at the optimal length.

During all steps, we continuously recorded the chordae tension, electrocardiogram, radial blood
pressure, 3D TEE, and surgical view (Figure 2, Video 1).

#### 175 II.4 <u>STATISTICAL ANALYSIS</u>

Baseline and demographic categorical variables were expressed as percentages, while quantitative
variables were expressed as medians (first and third quartiles, interquartile range).

178 Chordae tensions were expressed as mean ± standard deviation of peak-to-peak amplitude value 179 because they are the physical parameters representing the force that sustained the mitral valve. It is 180 therefore more relevant to know the extrema to not exceed the critical values. In addition, an average 181 measurement is achieved in real time to avoid noise of the sensors and of the electronic conditioning. 182 Categorical variables are presented as numbers with percentages.

The Wilcoxon paired test was used to compare chordal tension before and after optima correction. A
P-value <0.05 was considered to be significant. SPSS statistical software was used (IBM SPSS</li>
Statistics, version 24.0. Armonk, NY, USA).

# 187 III. <u>Results</u>

188 III.1 <u>CLINICAL RESULTS (ASIDE FROM INTRAOPERATIVE TENSION MEASUREMENTS)</u>

189 From 1 January 2016 to 1 September 2017, seven consecutive patients underwent transapical
190 neochordae implantation using the NeoChord DS 1000 (NeoChord Inc, Minneapolis, Minnesota,

191 USA). The characteristics and operative data of the seven patients are presented in

	All patients (n=7)	
Baseline clinical data		
Age	80	63 - 82
NYHA III/IV	5	(71)
Previous cardiac surgery	0	(0)
EuroSCORE 2	1,4	1,1 - 2,8
Baseline Echocardiographic Data		
LVEF (%)	66	57 - 68
SPAP (mmHg)	50	45 - 61
Severe MR	7	(100)
EROA (cm <sup>2</sup> )	0.51	0.47 - 0.58
Posterior prolapse	7	(100)
P2	5	(71)
P1/P2	1	(14)
P2/P3	1	(14)
volume LA (cm <sup>2</sup> )	33	27 - 36
LVDd (mm)	59,5	55 - 63
Operative data		
Number of neochordae implanted	3	3 - 4
Posterior localization	7	(100)
Operative time (min)	117	114 - 126
Autotransfusion volume (ml)	750	635 - 1025
Initial technical success	7	(100)
Conversion to conventional surgery	0	(0)
30-day outcomes		
ICU stay (days)	1	1 - 1,5
Hospital stay (days)	9	8 - 13
Discharge TTE		
None or mild MR	6	(86)
Moderate MR	1	(14)
Severe MR	0	(0)
Surgical revision for bleeding	0	(0)
Relevant Adverse Event		
Stroke	1	(14)
Perioperative infection	0	(0)
Transfusion	0	(0)

0 (0)

192

193 Table 2. The adjustment duration appears to have been reduced compared to the standard protocol.

194 The procedural success (defined as residual MR < Grade 2) was obtained in all patients.

195 The median per-procedural autologous autotransfusion volume (from cell-saver) was 750 196 ml (interquartile range 635-1025), and no patient required postoperative heterologous transfusion. The 197 mean hemoglobin level at discharge was  $10.2 \text{ g dl}^{-1}$ . No perioperative infection and no deaths were 198 observed after a mean follow-up of 10 months (1–18 months).

During initial follow-up, one patient had an early recurrence of severe MR due to a tear in a cleft-like indentation adjacent to the repaired P2 prolapse. He had a successful minimal-invasive MVr four months after the NeoChord procedure.

## 202 III.2 <u>TENSIONS DATA DURING LENGTH ADJUSTMENT PHASE</u>

Death

203 <u>Traction of the chorda implanted in the best location until best correction (Step 1)</u>: The amplitude of 204 force increased progressively until it reached a plateau value between 0.7 and 0.9 N (Figure 3). This 205 value was similar for all patients. Interestingly in all evaluations, we noticed an oscillation in the 206 amplitude of tension of approximately 13 % that followed the respiratory cycles (Figure 3).

<u>Traction of all chordae one after the other until homogeneous tension (Step 2)</u>: When we were setting the tension of the chordae, one at a time, aiming to achieve the same tension in all chordae, we observed a decrease in tension on chordae previously set in tension. At the end of Step 2, the tension initially measured on the first chorda was divided by the number of chordae set in tension. For example, when a tension of 0.8 N was observed with one chorda, it decreased to 0.2 to 0.3 N when four chordae are set in tension (Figure 4).

At the end of Step 2 the mean of the sum of the amplitude of tension in all patients was  $0.98 \pm 0.08$  N.

214 <u>A final adjustment under TEE control (Step 3)</u>: At this point of our protocol, some degree of

215 regurgitation usually persisted, thus requiring a final adjustment by synchronous traction on all of the

216 chordae, until optimal TEE results were achieved (no residual MR, vanishing of leaflet billowing and

217 homogeneous and high coaptation height).

218 While the initial mean of the sum of the amplitude of tension was  $0.98 \pm 0.08$  N before optimal 219 correction of MR, it decreased gradually to  $0.87 \pm .07$  N following the correction of regurgitation, 220 corresponding to a  $12 \pm 2$  % decrease (range: 10.3-14.1 %, p=,018) during step 3 (Figure 5).

#### 222 IV. Discussion

#### 223 IV.1 WHAT DID WE ALREADY KNOW?

The study of objective physical consequences of mitral valve repair remains limited by the central position of the valve in the heart and its complex structure. To date, our knowledge about chordae tension has previously relied entirely on animal experimental models.

In vitro studies have been published using an ovine experimental model<sup>4-8</sup>. The valvular tissue is 227 harvested en masse (ring, valve, and mitral sub-valvular apparatus). The valve and the papillary 228 229 muscle are attached to a left ventricular simulator, which can mimic the physiological hemodynamic regimen of the left ventricle. A miniature C-shaped force transducer is sutured in the middle of a 230 231 chorda. Interesting data were thus obtained, in particular concerning the differences in tension between marginal, struts, and basal chordae<sup>7</sup>. He and Lowers proposed an alternative method to 232 reduce interference between the movements of the chordae and the valve<sup>9</sup>. This consisted of 233 234 measuring the longitudinal tension of the chordae by an extrapolation of the measurement of the 235 transverse tension of the chordae fibers. A small sensor (elliptical AFP4, Microstrain Inc., Williston, 236 VT) is inserted within the same chordae. Both approaches remain highly experimental.

Few *in vivo* studies have been published using a porcine experimental model<sup>10</sup>. A dedicated in-line pre-mounted force transducer is inserted between the papillary muscle and the leaflet, and the strain gauge wire is exteriorized through the apex. This model is the one that most approaches our technique, and the measured tensions are close to the values reported here.

The transapical neochordae implantations allow direct measurements of the tension applied on the neochordae of a mitral valve in a human beating heart for the first time. These data, formerly inaccessible, are thus innovative.

Here we report data relevant to the function of the mitral subvalvular apparatus in a beating heart. Our tension measurements, recorded throughout the cardiac cycle (from prolapse to tethered chordae), led us to the following conclusions.

249

# THE TENSION APPLIED TO CHORDAE HOLDING ONE MITRAL SEGMENT IS RELATIVELY LOW (0,8 TO 1 NEWTON).

252 The stress-strain curves of a native chordae had been reported by Zuo, showing ultimate stress values around 35 MPa (equals to 35 N mm<sup>-2</sup>)<sup>11</sup>. In other words, a rupture of chordae should happen if a 253 traction is applied on a normal chorda of  $1 \text{ mm}^2$  and of a prolapsed area if the tension is 35-times 254 255 superior to the physiologic traction. This is highly unlikely to happen, meaning that the ruptures of the 256 chordae we encountered among our patients are likely a consequence of the fragility of the dystrophic 257 chordae. Moreover, we noticed that the forces applied to the subvalvular apparatus vary with 258 respiration. This might account for why some exceptional thoracic traumatism with the sudden 259 increase of the intra-thoracic pressure can lead to traumatic rupture of normal chordae and acute  $MR^{12,13}$ . 260

Gore-Tex<sup>®</sup> sutures (CV-4 or CV-5) are typically used as neochordae. The strength of these sutures is
 perfectly adapted as artificial chordae<sup>14</sup>.

The measures presented here are only valid for primary human chordae of 0.7 to 0.8 mm, and which are attached to the free edge of the leaflets. This is probably different for secondary chordae (basal chordae or strut chordae), which are thicker (1.2 to 1.3 mm) and are not involved in neochordae implantations. Our team has already reported the various functions of primary and secondary chordae, and it is important to distinguish their specific roles<sup>15</sup>.

- 268
- 269
- 270

#### 272 THE TENSION APPLIED TO CHORDAE HOLDING ONE MITRAL SEGMENT IS DIVIDED BY THE NUMBER

#### 273 OF CHORDAE SET IN TENSION.

274 Tension decreased as the number of chordae set in tension increased. This finding, although intuitively
275 expected, is physically confirmed here, with an almost strict mathematical correlation.

The spider's web-structure of the subvalvular apparatus, with numerous chordae and numerous implantation sites on the leaflets, is essential.

We could extrapolate that it is also necessary to use several chordae to fix a prolapse segment all along its free edge, moreover if it is a large prolapse segment. An increased number of chordae would likely make the repair less fragile and more durable.

281

# 282 THE TENSION OF THE SUBVALVULAR APPARATUS IS LOW WHEN THE CONTROL OF THE 283 REGURGITATION IS OPTIMAL

This is the most important finding of this study. The normal valve seems to work in a low-stress regimen. In other words, in a physiological state, the perfect systolic closure of the mitral valve with two coapted leaflets inside a 3D saddle shape annular plan corresponds to the situation with the lowest tension on the subvalvular apparatus.

288 Few physiological explanations can be suggested to explain this finding (Figure 6):

- As mentioned above, the total tension is divided by the number of chordae. After MR
   correction, the global tension is distributed between native and artificial chordae. Thus, the
   tension applied on neochordae (the only one that we can measure) is decreased.
- In the classical posterior prolapsed situation, the shape of the pathological leaflet formed a
   funnel, leading the blood flow toward the prolapsed area. Thus, the tension of the subvalvular
   apparatus of this zone is maximum.
- Because the two leaflets lean one against the other, the vertical vector of tension applied on the
   free edges (towards the left atrium) decrease and thus the measured tension on the primary
   chordae decrease.

The force of coaptation might increase. This hypothesis is supported by Falk et al who found that greater coaptation distance might contribute to a more durable result after repair<sup>16</sup>.

- A part of the tension is transferred laterally, to the secondary chordae (not measured here), to the annulus and to the heart walls. Nielsen et al., also found a significant reduction of the tension on the secondary chordae after mitral ring annuloplasty, without affecting the tension on the primary chordae<sup>17</sup>.
- The absence of regurgitation is associated with a normal intraventricular blood flow, with a
   normal clockwise vortex<sup>18</sup>. The ejection flow is then directed more toward the outflow tract
   and less to the mitral valve lowering the chordae tension.

This observation gives a plausible proposition of the natural history of the primary MR. As the valve starts to leak, the tension on the chordae increases, leading to a vicious cycle and an accelerated progression of the regurgitation. Whereas restoring a large surface of coaptation increases this keystone effect, and decreases the stress on the chordae, ensuring the long-term stability of the surgical repair, as usually reported in clinical settings<sup>19-20</sup>.

While not directly supported by this paper, it is likely that downsizing annuloplasty leads to a greater apposition of the two leading leaflets, with the keystone effect leading to a decreased tension on the chordae.

While we are approaching the limits of existing tools to explore mitral valve function and the consequences of repair techniques, we are continuing to develop innovative tools (numerical simulation, medical imaging, new sensors) to assess these hypotheses.

In conventional surgical MVr, the only existing tool available to evaluate immediate success of a repair is the saline test (also potentially paired with an ink-test). When neochordae are implanted the length adjustment on arrested hearts relies mainly on the habits of the surgeons. Dozens of tips and tricks have been proposed<sup>21</sup>.

A change of paradigm recently occurred with the introduction of novel techniques performed on beating hearts like transapical neochordae implantations allowing real-time adaptation of the repair thanks to an intraoperative TEE control. It is also possible to fix the length of the chordae exactly to the optimal position; neither too long nor too short. This represents a potential advantage of transapical neochordae implantation over traditional surgical techniques.

329 This feature has been exploited in this work: additionally, to a strict morphological assessment, the 330 plasty was optimized thanks to an objective parameter. Recent large cohort evaluating mid-term 331 results of NeoChord reported 10% of technical failure despite initial good echocardiographic results<sup>23</sup>. 332 We can assume that for 10% of cases TEE was not sensitive enough alone in guidance for NeoChord 333 placement and adjustment. We find that in optimal repairs chordae tension was minimum and 334 homogeneously spread on all chordae. We can thus hypothesize that measurement of chordae tension 335 could be use be useful in addition of TEE to insure optimal procedure. However, these beneficial 336 characteristics remains to be validated by additional studies.

Finally, although our measuring device was initially developed to assess chordae tension, its userevealed interesting features for chordae length adjustment:

Simplicity while avoiding iterative forceps manipulation on neochordae at cardiac apex,
 without increasing the operative time (operative time of our center with the use of the platform
 are similar with those reported by reference centers<sup>22</sup>). The set-up of the platform is simple.

- Precision thanks to a millimetric per-chorda or global adjustment.

- Safety thanks to permanent control of applied tension during the procedure avoiding excess
traction on the frail valvular tissue.

Thus even without tension assessment, the use of the device may also be beneficial for safety and precision.

347

#### 348 IV.4 <u>LIMITATIONS OF OUR STUDY</u>

The neochordae (ePTFE CV4 GORE-TEX<sup>®</sup> suture, Gore Medical) attached to the valve comes out 349 350 through the ventricle wall. The resistance of this passage way is not quantifiable, but it is probably 351 minimal. Relative to the resistance, the length and traction direction of the neochord (from the free 352 edge to the apex) are more important, which are different from those of the native chordae (from the 353 free edge to the papillary muscle) (Figure 7). Even if the Neochord procedure recommends the 354 introduction of the device as laterally as possible, we remain at some distance from the papillary muscles, and it is likely that the forces applied on the native chordae are slightly different 10,24. 355 356 Nevertheless, the global understanding of the sub-valvular function and its usefulness for assessment 357 of the optimal length during this procedure remains valid.

#### 358 IV.5 <u>Perspectives</u>

Our findings have helped generate new hypotheses about mitral valve function, as well as raisingmany as yet unanswered questions.

361 Concerning clinical perspective, the prognostic value of measured tension and its ability to improve 362 long-term success repair rate will be obtained thanks to the prospective follow-up of this cohort. In 363 our center, the platform is used for all implantations. We are currently pursuing the optimization of the 364 platform before its duplication to allow its assessment by other expert teams.

365 Concerning experimental perspectives, we reached some limits of our clinical model. *Ex-vivo* studies 366 are currently performed to compare the different technologies for chordae tension measurement as 367 well as measuring the impact of the differences in axis and length of neochordae implanted 368 transapically compared to the conventional implantation. Larger project involving multiple 369 technologies (medical imaging, numerical simulation) are required to better understand the changes in 370 structure and flow occurring during transapicale neochordae implantations. From a more global perspective, in conventional surgical repairs many different repairs techniques are continuously described, and multiple prosthetic devices are available (e.g. band, rigid annuloplasty ring, etc). Nonetheless, morphologic assessment tools failed to determine the best surgical options and the surgeon's expertise primes above objective concrete data.

New beating heart valve treatments have already grown in popularity as devices and quality of the 3D images improved. Percutaneous repair techniques have a single anatomical target, in opposition to the conventional surgery that often deals with multiple repair actions. Combo procedures appears promising. But as for conventional surgery, the role of annuloplasty to other techniques remains to be determined as well as the criteria for their implantations.

380 Original measuring tools allowing real-time intraoperative measurement (in addition to standard 381 morphologic evaluation) could increase our capacity to treat patients with customized techniques 382 guided by objective data during surgical and percutaneous interventions.

## 384 V. Conclusion

Transapical neochordae implantation conducted on a beating and functional heart, provides an excellent model for *in vivo* study of the tension forces on neochordae. This work brings original and useful data concerning the physiopathology of the sub-valvular apparatus: the tension applied on human chordae to sustain P2 segment of mitral posterior leaflet is low (1 Newton); this tension is divided by the number of chordae holding this segment; during MVr the lowest tension on all neochordae is observed when the control of the regurgitation is optimal. Such advances help to better understand MVr and, consequently, may play a role in the future treatment of MV disease.

- 392 Along with this objective assessment, the tool developed allows a safe precise and reproducible
- 393 procedure.

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# 464 VI. Figures and tables



467 Figure 1. The novel chordae tension measuring device allows the assessment of tension applied on neochordae during a
 468 transapical implantation. This pictures shows the platform after fixation of three neochordae. Each neochordae is attached
 469 to a piezo-resistive sensor thanks to a crocodile clip. The micrometer screw system allows pulling of each chordae one
 470 after the other or all simultaneously.







477 Figure 3. The amplitude of tension during Step 1 of the protocol: we started pulling the first chordae implanted in the
478 middle of the prolapse zone. During this traction the amplitude of tension increqsed until it reached a plateau value
479 between 0.7 and 0.9 N. The tension oscillated, and the oscillation corresponded to the mechanical ventilation.





#### 483 Figure 4. Schematic representation of the evolution of Amplitude Of Tension (AOT, in Newton) in chordae during

**Step 2:** While pulling the first chorda (yellow), the amplitude of tension was 0.79 N. The other chordae were then pulled

485 consecutively. At the end of this phase the initial tension was divided by the total number of chordae set in tension.





#### 497 Figure 6. Hypotheses explaining the change in chordae tension under normal and pathological states.

Three main hypotheses may participate to the reduction in total chordal tension occurring during the fine adjustment of chordae traction:The loss of "funnel effect" directing the flow towards the prolapse zone (left); The mutual abutment of the leaflets edge-to-edge with a "key-stone effect" (middle); and the return to normal clockwise intraventricular vortex with an ejection flow directed toward the outflow tract and no toward mitral leaflets (right).





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#### Figure 7. Difference in axis between native chordae and transapical neochordae.

504 In this case the apical neochordae insertion site was marked with a metal clip during the procedure. A postoperative CT-505 scan was performed allowing the visualization of the free edge of the posterior leaflet, the summit of the posterior papillary 506 muscle, and the apex. The angle of the native chordae (between the papillary muscle and the free edge of the leaflet) and 507 the neochordae (between the clip and the free edge of the leaflet) was 54°. The length of the native chordae was also 508 shorter than that of the neochordae.

Step 1	Traction of the chorda implanted in the best location until best correction
Step 2	Traction of all chordae one after the other until homogeneous tension
Step 3	Synchronous traction until optimal TEE correction
Step 4	Fixation of the chordae at the apex of the left ventricle at the optimal length

Table 1- Steps of the measurement protocol