## Chordae Tendineae Repair Techniques: A Literature Review

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## Abstract

Advancements in cardiac surgery have significantly transformed the management of mitral valve (MV) disease, with chordae tendineae repair techniques emerging as crucial for restoring heart function and preventing heart failure. Among these, the neochord procedure has gained increasing popularity as a primary repair approach. Many studies longitudinally followed patients who received neochord procedures to see their overall, long-term outcomes. In general, these studies found long-term outcomes to be positive, including low rates of disease recurrence and reoperation. Studies continue to demonstrate that the neochord procedure is effective at mitigating MV regurgitation while also improving overall heart function. Complications have been found to be rare for this procedure. These findings position the neochord procedure as a safe, effective, and increasingly preferred surgical option for appropriately selected patients.

While current evidence strongly supports the safety and efficacy of the neochord procedure, future research should focus on large-scale, randomized controlled trials comparing neochord to other minimally invasive and conventional mitral valve repair techniques. Additionally, further studies are needed to evaluate the long-term durability of neochord implants beyond the current follow-up periods, explore patient selection criteria, and optimize procedural techniques to enhance outcomes.

## Introduction

Progress in cardiac surgery has transformed the treatment of mitral valve (MV) disease, with chordae tendinae repair techniques playing a vital role in restoring heart function and preventing heart failure. Neochord is a primary repair technique that has grown in popularity. As a minimally invasive procedure, it replaces damaged chordae tendinae in MV repair. Techniques like neochord help prevent MV regurgitation (MR). MR occurs when the MV, between the left atrium and ventricle, does not close tightly. In this case, improper sealing can cause blood to flow back into the atrium, causing the heart to work harder. MR is caused by myocardial infarction, rheumatic heart disease, and MV prolapse. With its growing popularity for patients with severe MV disease, it is important to understand the overall outcomes, risks, and benefits of the neochord procedure. In this review, we investigate the current literature pertaining to the outcomes of the neochord procedure, as well as other similar valve repair surgical techniques that may have advantages and/or disadvantages compared to the neochord procedure.

## **Materials and Methods**

This study consists of a systematic review that addresses the question: *In patients with mitral valve regurgitation, how do clinical outcomes, complication rates, and long-term stability of the neochord procedure compare to other conventional mitral valve repair techniques?* An encompassing search of existing systematic reviews and randomized control trials (RCT) involving databases such as Pubmed was conducted. Data from the randomized controlled trials were selected including the study's design plan, participant demographics, outcomes, and reasoning for findings. Using all of the results of the trials and the information gathered, this literature search allowed for a comprehensive outlook on the advantages and disadvantages of neochord.

#### Search strategies

An electronic search without time or language restrictions was conducted using Pubmed, Google Scholar, and other published articles. The reference lists of included studies and relevant reviews were also searched for other potential studies. The detailed search strategies were as follows: #1 neochord, #2 mitral valve, #3 harpoon, #4 chordae transposition, #5 MitralStitch.

#### Discussion

## Neochord

#### Outcomes

Chordae tendinae help maintain one-way flow in the heart and can be damaged by stresses like myocardial infarction. Neochord is a chordae tendinae repair technique that prevents MR. Several studies investigating a variety of outcome measures after the neochord procedure suggest it is a safe and effective procedure. Many studies longitudinally followed patients who received neochord procedures to see their overall, long-term outcomes. In general, these studies found long-term outcomes to be positive, including low rates of disease recurrence and reoperation. In a 2020 observational study with 746 patients, the rate of reoperation with death was 4.2% at 20 years. Moderate to severe MR occurrence was 14.1% after 20 years. (David et al., 2019) Another study involving 29 patients found that patients classified as New York Heart Association (NYHA) functional class 3 or 4 before the neochord procedure ended up in classes 1-2, 23.7 ± 12.8 months after the surgery. (Cimen et al., 2006) Another study involving 111 patients found the five-year success rate of the neochord procedure, defined by survival, was  $98.2 \pm 1.8\%$ , and freedom from grade 1+ regurgitation rate was 97.2 ± 2.8%. (Chiappini et al., 2006) These studies demonstrate that the neochord procedure effectively accomplishes high rates of success, low rates of reoperation, and effective management of MR. More long-term research outcomes looking at mortality and morbidity would help to better characterize the ultimate impact of neochord procedures on patient-centered outcomes.

Few studies also looked at other surrogate outcomes after neochord surgery. One observational study investigated the hemodynamic profile at rest and at peak exercise of 24 patients who had undergone neochord surgery by measuring pressure gradients in the heart via echocardiography. This study found that patients had good hemodynamic profiles and also good exercise capacity. (Ragnarsson et al., 2017)

Another observational study with 66 patients assessed the outcomes of patients after neochord surgery by measuring left ventricular geometry and found that the LV end-diastolic volume had a 14% reduction, showing improved cardiac efficiency and a decrease in volume overload. (Colli et al., 2018) Other studies also measured and found reduced chordae tension after neochord, indicating a positive outcome. In one of these studies involving seven patients, chordae tendinae tensions were measured with transesophageal echocardiography (TEE) before and after neochord. Before neochord, chordae tendinae tensions ranged from 0.7-0.9N and decreased on average by 12% after neochord. (Grinberg et al., 2019) Another study measuring these chordae tendinae tensions, but in sheep hearts, found that the neochord procedures could reduce tension on the chordae to the baseline value of  $0.21 \pm 0.14$  N for those sheep. (Zhu et al., 2024) Overall, these measurements show that neochords reduce overall tension in the valve and the chordae, allowing for better efficiency and durability in the procedure and for long-term outcomes. These studies continue to demonstrate that the neochord procedure is effective at mitigating MV regurgitation while also improving overall heart function.

## **Complications of Neochord**

In a 2020 case report, two patients had undergone a neochord procedure and later developed MV dysfunction due to valve remodeling, which led to MV regurgitation. (Moore et al., 2020) Another case report showed a unique case of severe hemolytic anemia and acute renal failure after having a neochord surgery, caused by a mild MR striking the annuloplasty ring, leading to complications. (Li & Duan, 2021) Finally, a 2022 case report shows a case of intraoperative acute procedural failure during transapical off-pump MV repair using neochordae implantation due to a loss of tension secondary to leaflet-free edge tear. This led to moderate to severe MR. However, a second neochord implantation was done with a transapical access, leading to improvement, and the patient was discharged with mild regurgitation and a normal left ventricular ejection fraction. (D'Onofrio et al., 2021) The authors note that these complications were rare and that future ways to mitigate complication risk can include the use of different access points on a case-by-case basis if there are previous myocardial insults and prior prediction of possible outcomes before the surgery.

#### **Animal studies**

Many promising results have been seen not only in humans but in animals and computational models as well. These tests can prove that neochord can balance stress and improve overall valve function. A 2019 intervention study used porcine MVs in a 3D-printed heart simulator to test out polyvinyl alcohol nanofiber (PVAN) neochords, which are known to prevent MR caused by a posterior leaflet prolapse. This study found that PVAN neochords allowed the forces on the chordae tendineae to return to baseline levels, suggesting that PVAN neochords are effective, minimally invasive, and put less stress on chordae tendinae. (Paulsen et al., 2020) A different study used 8 porcine MVs to evaluate the importance of leaflet preservation in different surgical strategies to fix MV prolapse for chordal rupture. Neochordoplasty led to a significant decrease in regurgitant volumes, as well as returned coaptation length to baseline levels. (Padala et al., 2009) Also, a recent 2024 study used sheep hearts to evaluate

different MV repair techniques, including neochord, non-sectional leaflet remodeling, and triangular resection. Among these methods, neochord procedures have done the best at reducing peak chordal forces, bringing them closer to baseline values. (Zhu et al., 2024) In summary, the evidence of successful neochord procedures is reinforced in the animal studies we present, and future studies should take advantage of studying novel techniques and methods in animal hearts to find ways to improve MV surgical management in humans.

## **Pediatrics**

Studies suggest the neochord procedure is effective for pediatric patients, with favorable outcomes in short and long-term settings. An observational study following 21 children to evaluate the outcomes of pediatric atrioventricular valve repair with artificial chordae found that there was no deaths or valve restrictions, the only complications to be moderate tricuspid insufficiency and moderate mitral insufficiency experienced by 2 patients each at the last follow-up which was around 28 ± 18 months, two patients had to go under reoperation(for mitral ring dehiscence recurring MV insufficiency). (Boon et al., 2007) Another observational study with 78 pediatric patients examined the long-term results of neochord MV repair with ePTFE. Of the 78, there was one in-hospital death due to left ventricular dysfunction, severe aortic regurgitation, and moderate MR, and three experienced moderate regurgitation. Despite this, there was an overall high rate of freedom of reoperation, which was 90.4% at 10 years. Additionally, echocardiography confirmed good heart and valve function at the 1-15 year mark. (Oda et al., 2013) A different observational study followed 15 infants to find the long-term effects of MV repair, focusing on durability with ePTFE sutures in infants with severe MR. There were no operative or hospital deaths, and there was a 10-year survival rate of 89%. (Murashita et al., 2012) Overall, the short-term and long-term outcomes of severe MR repair in infants with the neochord procedure are satisfactory. The procedure is also durable and safe, and the elevated risk of long-term adverse effects is low.

## Harpoon

Another technique that has not been widely used but may gain popularity in the future is the Harpoon technique, which is similar to neochord with the exception that it is placed with automated, catheter-based deployment. Studies have shown that Harpoon is a reasonable option for minimally invasive techniques, offering specific advantages. Neochord, using a posterior access site, had lower leaflet stress and improved coaptation during systole, which made it effective for Barlow's disease, a form of MV prolapse that requires precise and durable support to properly close valves during systole. On the other hand, the Harpoon technique utilizes anterior access, which allows it to perform better in some cases, but studies found greater variability in chordae tension forces, making it unstable. Moreover, anterior access methods increase the risk of suture-leaflet contact, so incision sites must be precise. (Micco et al., 2022) Another observational study was conducted to assess the effectiveness of the Harpoon system for minimally invasive MV repair in patients with severe MR secondary to posterior leaflet prolapse. Similar to neochord, the Harpoon method helps avoid cardiopulmonary bypass and cardioplegia. Even though the stabilization of the device was hard due to

the extensive leaflet flail, the result was successful as it yielded MV competence after attaching the implanted chordae. (Gerosa et al., 2017) Overall, Harpoon has the potential for minimally invasive techniques for MV repairs, but specific care in the variability in tension and stabilization will be needed in order to ensure positive outcomes.

# **Chordae Transposition**

Overall, studies have shown that chordae transposition, a strategy that takes chordae from places of less need to greater need, can have some promising results and be an effective strategy. The procedure of chordae transposition is used to include isolated mitral insufficiency, mitral stenosis, tricuspid valve repair, and ruptures in MV chordae transposition. In an interventional study featured, 29 patients, 21 with isolated mitral insufficiency and 8 with both mitral insufficiency and stenosis. Before the surgery, 12 patients were in the NYHA Functional Class II, 16 were in Class III, and one child was in Class IV. After the surgery, a follow-up showed that 24 out of 26 patients were in Class I, which shows improvement. Additionally, 17 patients had no heart murmurs, 5 had mild, and 4 had moderate murmurs. However, two in-hospital deaths occurred due to severe arrhythmia and respiratory failure. (Lessana et al., 1988) A case report for a 48-year-old male reports that one month after the procedure, echocardiography showed only mild tricuspid regurgitation with no transvalvular gradient, cardiomegaly improved, with a reduced cardiothoracic ratio of 55%; and after a year, the patient remained well with a strong valve. (Takamatsu et al., 2018) Finally, an interventional study that used computational modeling found that after the repair, the stress was reduced from 0.61 MPa to 0.37 MPa, showing ideal results, and leaflet coaptation was restored (with a coaptation length of 6.37 mm). Additionally, the anterior leaflet coaptation angle decreased to 4.0° post-repair(8.8° is normal), while the posterior leaflet angle increased to 43.3°(31.5° is normal), which helped improve leaflet coaptation and valve function and could restore normal geometry. Stabilizing the posterior leaflet more than the anterior leaflet can help maintain coaptation and support chordae tendinae. Even though the angles differ from normal leaflet angles, they are still acceptable as they restore normal MV function and redistribute stress. (Jeong et al., 2023)Overall, these studies show that chordae transposition is a good repair strategy; it is durable, and it can also avoid some cases of valve replacement entirely.

# MitralStitch

The novel method known as MitralStitch could have probable benefits in the future in regards to safety. MitralStich is a method like neochord, but it uses sutures rather than artificial chords. In these procedures, MitralStitch was used to fix MR. Also, testing safety, effectiveness, and outcomes. A 2022 case report with 10 participants found that one year after the procedure, the results were mixed, with 4 out of 10 patients having some MR. (Saccocci & Colli, 2022) A case report in the same year using 10 patients found that after Mitralstitch, 5 patients had little to no MR, and 5 patients had mild MR; after a 1-month follow-up, all patients were safe, and an efficacy endpoint was completed. (Wang et al., 2022) Another case report involving just 1 patient showed that after the surgery was complete, a TEE showed that the 78-year-old man had no MR, the MV had neocoaptation length, and positioning was good. (Wang et al., 2018)

Overall, the surgery had mixed short-term results and was easily reproducible, but further studies should test more long-term effects, efficiency, and refinement could be needed for better outcomes and in order to allow it to be used widely.

## Conclusion

In this review, we investigated the current literature on the outcomes of the neochord procedure and also looked at complications, alternative surgical methods, the evidence in animal studies, and use in pediatric populations. The neochord is safe and effective in both adult and pediatric populations; it will continue to be a first-line surgical management option for patients with the appropriate indications. In the future, more research can focus on optimizing neochord strategies, developing further patient-specific usages, and studying further long-term outcomes to advance neochord and patient care.

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