

Functional Outcome and Complications of Wide Excision of the Distal End of Radius and Reconstruction with Ulnar Transposition in Campanacci Grade III Giant Cell Tumor: A Longitudinal Study of 14 Cases in South India

S Sai Krishna¹, Dijoe Davis², K Ajith John², Dominic K Puthoor²

Learning Point of the Article:

Wide excision followed by ipsilateral ulnar transposition serves as a robust biological reconstruction for Campanacci Grade III giant cell tumors of the distal radius, providing stable wrist arthrodesis and preserving forearm rotation without the donor-site morbidity associated with distant autografts.

Abstract

Introduction: Giant cell tumor (GCT) of the bone is an intermediate, locally aggressive neoplasm with a high incidence in the Indian subcontinent. The distal radius is a frequent site, often presenting with extensive cortical destruction (Campanacci Grade III), necessitating en bloc resection. Reconstructing the resulting defect remains a challenge in orthopedic oncology. This study evaluates the functional and oncological outcomes of ulnar transposition after radial resection.

Material and Methods: A cohort of 14 patients with Grade III GCT of the distal radius underwent en bloc resection and reconstruction with ipsilateral ulnar transposition between 2015 and 2023. Functional outcomes were assessed using the Musculoskeletal Tumor Society (MSTS) score, grip strength measurements, and range of motion (supination and pronation). Radiological union times at the ulno-radial and ulno-carpal junctions were recorded.

Results: In our study, we have a total study sample of 14 cases. Both males and females have equal preponderance for GCT of bone. Mean age was calculated to be 39 ± 13.3 years. Mean of the VIII MSTS score was 26.21 ± 1.67 . Majority of the patients (78%) have no pain post-operatively. Mean time taken for ulno-radial union was 6.5 months and for ulnocarpal union was 5.7 months. There is a strong agreement of Hand Grip Strength score between affected and unaffected hand Cronbach's alpha +0.961. Mean of supination was 83.57 ± 8.419 (range 70–90°). Mean pronation was 75 ± 9.405 (range from 60 to 90°). The mean of actual grip strength of the affected hand is 16.15 Kg as compared to 31.78 kg for the opposite side. In 64% of cases were not having any post-operative complications associated with surgery. There were only 4 cases (28.5%) of implant failure associated with non-union. One case (7%) of recurrence was noted.

Conclusion: Ulnar transposition is an effective, technically accessible biological reconstruction that avoids the complications of fibular grafting or prosthetics. It provides a stable, painless wrist while preserving significant forearm rotation, making it a viable option for aggressive distal radius tumors.

Keywords: Giant Cell Tumor, Distal Radius, Ulnar Transposition, En-bloc Resection, Wrist Arthrodesis, Biological Reconstruction, Orthopaedic Oncology

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Introduction

The clinical entity known as giant cell tumor (GCT) of the bone has long fascinated and challenged the orthopedic community since its initial description by Sir Astley Cooper in 1818. Historically, the understanding of GCT has progressed from a poorly defined lytic lesion to a well-characterized neoplasm under the World Health Organization (WHO) classification, which identifies it as an osteoclastic giant cell-rich tumor of intermediate nature [1]. While fundamentally benign, its biological behavior is remarkably unpredictable, characterized by local aggressiveness, significant osteolytic potential, and a documented, albeit rare, capacity for pulmonary metastasis, occurring in 1–9% of cases [2].

The epidemiological landscape of GCT reveals a striking geographic disparity. In Western populations, GCT accounts for approximately 5% of all primary bone tumors. In contrast, reports from East Asia and specifically Southern India indicate a much higher prevalence, with some series suggesting it constitutes up to 20% of all primary bone neoplasms [3]. The distal end of the radius is the third most common location for GCT, following the distal femur and the proximal tibia [4]. This site is particularly significant due to the complex anatomy of the radiocarpal joint and the high functional demands placed on the wrist in young adults, who are the most frequently affected demographic [5].

The pathogenesis of GCT is rooted in a unique cellular environment. The lesion is composed of mononuclear spindle-like stromal cells, which are considered the neoplastic component, and a massive secondary population of multinucleated osteoclast-like giant cells. The recruitment and activation of these giant cells are driven by the overexpression of the receptor activator of nuclear factor kappa-B ligand (RANKL) by the stromal cells. This RANK-RANKL interaction triggers intense osteoclastic resorption, leading to the rapid destruction of the subchondral bone and cortical expansion [6]. In Campanacci Grade III lesions, this process leads to the complete breach of the bony cortex and extension into the surrounding soft tissues, significantly increasing the risk of local recurrence if the primary surgical management is inadequate.

Surgical management remains the cornerstone of treatment for GCT. For less aggressive Grade I and II lesions, extended curettage with the use of local adjuvants such as phenol, liquid nitrogen, or bone cement has shown acceptable results in reducing recurrence [7]. However, for Grade III lesions,

where the integrity of the joint surface and the cortical shell is lost, intralesional curettage carries a recurrence risk as high as 65% [8]. En bloc resection offers superior oncological control but leaves a massive segment of missing bone at the distal radius. The subsequent reconstruction must address three goals: Providing a stable support for the hand, preserving as much forearm and hand function as possible, and ensuring long-term biological durability [9].

Various reconstructive options have been described in the literature. Osteoarticular allografts provide an anatomical match but are fraught with risks of infection, non-union, and late articular collapse due to the lack of biological vitality. Vascularized fibular autografts are considered a gold standard for biological reconstruction but involve complex microvascular surgery and significant donor-site morbidity, including common peroneal nerve injury and ankle instability [10]. Custom-made prostheses offer the advantage of early mobilization but are expensive and carry the risk of mechanical loosening and infection over the long term [11].

Ulnar translocation, originally described by Seradge in 1982, offers a pragmatic and biologically sound alternative [12]. Using the ipsilateral ulna to bridge the defect, the surgeon utilizes a local source of autologous bone that maintains its



Figure 1: Surgical steps of ulnar transposition.



Figure 2: Patient underwent en bloc resection and ulnar translocation with a good functional outcome. This patient has good forearm movements of 90° of pronation and 80° of supination and hand grip strength of the affected side of 75% of the opposite side. Patient has a musculoskeletal tumor society score of 28, suggestive patient is having good functional upper extremity in terms of function, hand position, manual dexterity, emotional acceptance, and lifting abilities.

vascularity through its native muscle attachments. This procedure eliminates the need for a secondary surgical site and avoids the complications associated with microvascular anastomosis. Despite its advantages, the procedure results in wrist arthrodesis, which necessitates a careful evaluation of the functional trade-offs. This study aims to provide a comprehensive analysis of the functional outcomes and associated complications of wide excision and ulnar transposition in a cohort of 14 patients treated at a tertiary care center in South India, contributing to the limited body of evidence regarding this specific reconstructive technique.

Materials and Methods

Study design and population

This research was structured as a cohort study conducted at the Department of Orthopaedics at Amala Institute of Medical Sciences (AIMS), a prominent tertiary care hospital in Thrissur, Kerala. The study focused on patients diagnosed with Campanacci Grade III GCT of the distal radius between the years 2015 and 2023. The inclusion criteria were stringent, requiring histological confirmation of GCT and radiological

evidence of Grade III disease according to the Campanacci classification. Patients who underwent en bloc resection and ipsilateral ulnar translocation were followed longitudinally to assess functional and radiological outcomes. The study was initiated following approval from the Institutional Ethics Committee (Ref No: 17/EC/22/AIMS-68).

Pre-operative staging

Pre-operative evaluation began with a detailed clinical history and physical examination, focusing on pain duration, the presence of a palpable mass, and the degree of wrist joint impairment. Standard radiographic views (AP and Lateral) were obtained to assess the lytic lesion, cortical destruction, and potential pathological fractures. Advanced imaging played a crucial role in staging. Computed tomography (CT) scans were utilized to classify the tumor's intraosseous or extraosseous extension. They were divided into the following classes based on the results of the CT scan. Intraosseous Class I tumors lacked cortical fractures. Extraosseous lesions with cortical fractures limited to a single surface and not extending beyond one-third of the circumference of the bone were classified as Class II tumors. Tumors classified as Class III consisted of extraosseous lesions that had penetrated the cortex on many surfaces or covered more than one-third of the bone's circumference [13].

Magnetic resonance imaging (MRI) was indispensable for determining the extent of soft tissue involvement and the proximity of the tumor to critical structures such as the median nerve and the radial artery. The level of radial resection was meticulously planned on MRI, ensuring a 1 cm margin of healthy bone proximal to the lesion.

Surgical technique

The procedures were performed by experienced orthopedic surgeons under general anesthesia. A pneumatic tourniquet was

Table 1: Age and gender distribution of the GCT cohort

Age group	Frequency	Percentage	Male	Female
21–30	4	28.6	2	2
31–40	5	35.7	3	2
41–50	2	14.3	1	1
51–60	2	14.3	1	1
61–70	1	7.1	0	1
Total	14	100	7	7

GCT: Giant cell tumor





Figure 3: Case that has underwent en bloc resection of the distal end of the radius and ulnar transposition. The post-operative period was uneventful. She was followed up and serial X-rays showed union of ulno-radial and ulno-carpal junction at the 4th month and 6th month, respectively. Her musculoskeletal tumor society score was 26, suggesting she was able to do daily activities without much limitations. A year later, when she came in for follow-up, an X-ray was taken which showed a broken implant.

used for all cases to ensure a bloodless field, which is essential for identifying the thin, expanded cortex and the margins of the tumor. The approach utilized a dorsal longitudinal incision. The superficial branch of the radial nerve was identified and carefully protected. The forearm musculature was dissected to reveal the tumor, with the second and third extensor compartments being the primary access points.

The en bloc resection involved the removal of the distal radius with a margin of 2 cm (based on T1W MRI image), the tumor, and a cuff of healthy soft tissue. Proximal to the tumor, a transverse osteotomy was performed based on the pre-operative MRI planning. Following the excision of the radius, the focus shifted to the ulna. The distal third of the ulna was exposed, taking care to preserve the muscular attachments of the pronator quadratus and the extensor carpi ulnaris to maintain the blood supply to the bone. An ulnar osteotomy was performed at a level corresponding to the length of the radial defect.

The ulnar segment was then translocated laterally into the position previously occupied by the distal radius. Arthrodesis of the wrist was performed by removing the articular surface of the

carpal bones (primarily the lunate and scaphoid). The translocated ulna was then stabilized to the proximal radius and the distal carpal bones using a dynamic compression plate (DCP). The plate was carefully contoured to accommodate the shift from the radial to the ulnar axis. The forearm was fixed in a mid-prone position to maximize its functional utility for daily activities. Before closure, the range of forearm rotation was assessed to ensure that the hardware did not mechanically interfere with pronation and supination (Fig. 1).

Post-operative protocol

The post-operative regimen involved initial immobilization in a volar splint until wound healing and suture removal. This was followed by a below-elbow cast for a period of approximately 2 months. Early finger movements and elbow exercises were encouraged from the 1st post-operative day. Graded forearm rotation exercises were initiated once early radiological signs of union were observed at the ulno-radial junction. Debusumab, chemotherapy, and radiation are not used pre or post-operative. Debusumab is usually reserved for soft tissue recurrence.

Outcome measures

The primary outcome was the functional assessment using the Musculoskeletal Tumor Society (MSTS) score.

This scoring system evaluates six parameters: Pain, function, emotional acceptance, hand positioning, manual dexterity, and lifting ability. Each parameter is assigned a score from 0 to 5, providing a maximum possible score of 30.

Secondary outcomes included:

1. Grip strength: Measured using a Jamar hand dynamometer at the final follow-up and compared to the contralateral unaffected hand
2. Range of motion: Pronation and supination were measured in degrees using a standard goniometer

Table 2: Functional assessment summary (range of motion and MSTS)

Parameter	Mean value	Standard deviation	Range (Min–Max)
MSTS score	26.21	1.672	23–28
Supination (degree)	83.57	8.419	70–90
Pronation (degree)	75	9.405	60–90
Grip strength (affected)	16.15 kg	1.42	10–20 kg
Grip strength (contralateral)	31.78 kg	2.5	28–37 kg

MSTS: Musculoskeletal tumor society



Table 3: Radiological union times and complications

Outcome metric	Mean time (Months)	Frequency/Percentage
Ulno-carpal union	5.7 months	100 Union
Ulno-radial union	6.5 months	71.4 Union
Non-union (ulno-radial)	N/A	4 cases (28.6)
Implant failure	N/A	4 cases (28.6)
Local recurrence	N/A	1 case (7.1)
Pain-free status (post-op)	N/A	11 cases (78.6)

3. Radiological union: The time to union at both the ulno-carpal and ulno-radial junctions was determined by serial radiographs

4. Complications: Surveillance for implant failure, non-union, local recurrence, and wound-related issues.

Statistical analysis was conducted using the Statistical Package for the Social Sciences version 23. Results for continuous measurements are presented as mean \pm standard deviation, while categorical data are presented as frequencies and percentages.

Results

The study cohort consisted of 14 patients, with an equal gender distribution of 7 males (50%) and 7 females (50%). This finding differs slightly from some international series that suggest a female preponderance for GCT, yet it reflects the diverse clinical presentation in the South Indian population. The mean age of the patients was 39.0 ± 13.3 years, with the youngest patient being 20 and the oldest 64. The most common age group affected was 31–40 years, accounting for 35.7% of the cases (Table 1).

Clinical presentation and staging

The majority of patients (78.6%) presented with the classic combination of pain and swelling at the distal end of the radius. One patient (7.1%) presented with a pathological fracture following minor trauma, which is a documented indicator of the aggressive nature of Grade III tumors. Pre-operative physical examination revealed palpable masses and tenderness in 85.7% of cases, with a significant reduction in wrist range of motion due to pain in 100% of the cohort. X-ray findings consistently demonstrated eccentric lytic lesions with cortical destruction, typical of Campanacci Grade III lesions.

Regarding co-morbidities, 64.3% of the patients were healthy. The remaining 35.7% presented with controlled type 2 diabetes mellitus, hypertension, or hypothyroidism. These systemic

conditions did not preclude surgical intervention but were considered in the post-operative healing phase.

Functional outcome: MSTS Score

The functional evaluation performed at the final follow-up yielded highly encouraging results. The mean MSTS score for the cohort was 26.21 ± 1.67 , with individual scores ranging from 23 to 28. A majority of the patients (71.4%) achieved scores of 26 or higher, which signifies an excellent functional outcome according to the Enneking criteria (Table 2).

The individual parameters of the MSTS score provided deeper insight into the patient's quality of life. Pain assessment revealed that 78.6% of patients had no pain post-operatively, while 21.4% experienced only mild, intermittent pain. Emotional acceptance was high, as the procedure allowed patients to return to their baseline activities and occupations. Manual dexterity and hand positioning scores indicated that despite the loss of wrist mobility, patients effectively compensated using their finger and forearm mobility.

Forearm rotation and range of motion

One of the primary advantages of ulnar translocation is the preservation of forearm rotation, which is often sacrificed in other types of wrist fusion. In this study, the mean supination achieved was 83.57 ± 8.42 and the mean pronation was 75.00 ± 9.41 . Over 57% of the patients regained full supination of 90° , and 64.2% achieved pronation of 80° or more. This preservation of rotation is critical for tasks such as using a keyboard, turning a doorknob, or holding a plate, which significantly impact the patient's functional independence.

Grip strength comparison

The hand grip strength of the affected side was compared with that of the contralateral normal hand using a dynamometer. The mean actual grip strength of the affected hand was 16.15 kg, whereas the unaffected side measured 31.78 kg. Thus, the affected hand retained approximately 51% of its normal strength. While this reduction is statistically significant, it was clinically tolerated by all patients. A strong correlation between the strength of the affected and unaffected hands was observed (Cronbach's alpha +0.961), suggesting that the surgical reconstruction provided a consistent and predictable functional result.

Radiological union

The time to achieve biological union is a critical metric for any reconstructive procedure. In our study, the mean time for

radiological union at the ulno-carpal junction was 5.7 months (ranging from 4 to 8 months). The ulno-radial junction, which represents the more proximal bone-to-bone interface, took slightly longer to unite, with a mean time of 6.5 months (ranging from 5 to 8 months for those that united) (Table 3).

Complications and recurrence

Despite the high functional scores, the procedure was not without complications. We observed non-union at the ulno-radial junction in 4 cases (28.6%). In all four instances, the persistent lack of union led to fatigue failure of the implant, necessitating revision surgery with bone grafting and replacement of the DCP. One case (7.1%) of local soft tissue recurrence was noted during the follow-up period. This recurrence was managed with the administration of debusumab, a monoclonal antibody against RANKL, which successfully downstaged the lesion to allow for a second excision with clear margins. There were no reports of surgical site infections, skin necrosis, or major neurovascular complications.

Discussion

GCT is a benign tumor but locally aggressive in nature. GCTs represent 4–10% of all primary bone tumors and 15–20% of benign bone tumors [14]. However, the report shows a higher incidence of GCT in the Indian and Chinese population as high as 20% in the Asian population [15]. In our study, the sample size was 14 patients which was comparable to Puri et al. In their study, the sample size was 14, and all patients were selected belonging to Campanacci grade III and underwent en bloc excision of the distal radius and ulnar transposition [16]. The problem of reconstructing following distal radius excision for GCT is due to multiple factors, including the high functional demands on the hand, restricted soft tissue around the area, close proximity to nearby nerves and tendons, young age, and comparatively long life expectancy of these patients. With advantages involving better functional outcomes, preserved vascularity, and the removal of donor site morbidity, en bloc excision with ulna translocation has established itself as a routine treatment option for GCT of the lower end of the radius. Microvascular expertise is not necessary for ipsilateral ulnar transfer following radial resection in comparison to free vascularized fibular grafting. GCT is a benign tumor but locally aggressive in nature. GCTs represent 4–10% of all primary bone tumors and 15–20% of benign bone tumors [14]. However, the report shows a higher incidence of GCT in Indian and Chinese populations as high as 20% in the Asian population [15]. In our study, the sample size was 14 patients which was comparable to

Puri et al. in their study, also the sample size was 14, all patients were selected belonging to Campanacci grade III and underwent en bloc excision of the distal radius and ulnar transposition [16]. The problem of reconstructing following distal radius excision for GCT is due to multiple factors, including the high functional demands on the hand, restricted soft tissue around the area, close proximity to nearby nerves and tendons, young age, and comparatively long life expectancy of these patients. With advantages involving better functional outcomes, preserved vascularity, and the removal of donor site morbidity, en bloc excision with ulna translocation has established itself as a routine treatment option for GCT of the lower end of the radius. Microvascular expertise is not necessary for ipsilateral ulnar transfer following radial resection in comparison to free vascularized fibular grafting.

The chief presenting symptoms include dull aching pain, localized swelling at the distal end of the radius lasting for few months. All the selected patients were healthy and none of them had any metastasis. One of the patients presented to us initially with a pathological fracture, Campanacci grade III GCT, distal end of radius. In our study, both males and females have equal preponderance with 50%. On the contrary, Campanacci et al. noted a higher predilection for female gender [4]. The age group of patients with GCT presented to us was commonly in their 2nd and 3rd decade, and as the age progresses, the incidence of GCT was declining. This was comparable to that of Puri et al. in which the common age group GCT presented was between 20 and 40 years, and as the age advances, the incidence of GCT of bone decreases [16]. Mean age of GCT in our study group was 39 years \pm 13.3. The predominant age group was between 20 and 40 years (64% cases, i.e., 9 patients). The mean time taken for ulno-carpal union was 5.7 months, and the mean time taken for ulnoradial union was 6.5 months. Puri et al. [16] in their study show that the ulno-carpal junction, the mean time to radiological union was 4.4 months (3.0–7.0), while at the ulno-radial junction, it was 4.9 months (3.0–8.0). Salunke et al. in their study showed that the mean bone union time at the ulna to carpal junction was 4.5 (4–6) months and the ulna to radius junction was 6.5 (5–8) months which was comparable to our study [17].

There is a strong agreement of Hand Grip Strength score between affected and unaffected hand Cronbach's alpha +0.961. Mean of hand grip strength is 51.28% compare to the opposite side (range from 34% to 75%). The mean of actual grip strength of the affected hand is 16.15 kg as compared to 31.78 kg for the normal side in our study. 69% of cases had a good hand grip on the affected side, showing a good functional outcome. This result was comparable to that of Johan et al., in their study which shows that the hand grip strength mean was 55%



compare to the opposite side [18]. Vyas et al., in their study, a retrospective analysis of GCT lower end of the radius treated with en bloc resection and translocation of ulna, showed that the mean of affected side hand grip strength was 29 kg compared to 41 kg for the unaffected side [19]. In our study, the mean of MSTS score is 26 and ranges from 23 to 28. This was comparable to Johan et al. Their study also showed a mean MSTS score of 25 and Puri et al. showed a mean MSTS score of 26 [18]. This MSTS score shows that there is a good functional upper extremity. Pain assessment was done with the MSTS score which shows 11 patients out of 14 (>78.3%) with no pain post-operatively and 3 patients (21.7%) with mild pain.

Mean value of supination is 83.57 and standard deviation is 8.419. Mean of pronation is 75 and standard deviation is 9.405. Majority of patients have supination of 90° (57.1%) which indicates complete regain of supination. Post-operatively, 64.2% of patients have regained pronation more than or equal to 80° which indicates a good functional outcome. This value was comparable to Johan et al., mean of forearm movement 82.66° of supination and 81.54° of pronation [18]. McLean et al., in their study, used the ulnar translocation and reconstruction technique for Campanacci grade III GCT of the distal radius, showing an average range of motion of the forearm- 80° of pronation and 70° of supination [20]. We had 4 non-union at the ulno-radial junction, resulting in all 4 implant failures as a major post-operative complication. Those patients were again operated and the previous implant was removed. Then, bone grafting was done at the ulno-radial junction, followed by DCP, which was used to fix the ulno-radial junction. Bhan and Biyani, in their study, also show that there are delayed unions at the ulno-radial junction and require further procedure involving bone grafting at the non-union site. One of the cases presented with soft tissue recurrence at the surgical site post-operatively [21]. This patient was managed with denuzumab. As the tumor size shrank with the denuzumab, tumor was excised with the surrounding soft tissue. Puri et al., in their study, also found 3 soft tissue recurrences of GCT without any bony involvement [16]. There was no surgical site infection, skin necrosis, or wound dehiscence. After the reconstruction, the majority of patients, including those who experienced issues, were able to return to their regular activities and previous jobs.

Case showing patient underwent en bloc resection and ulnar translocation with a good functional outcome. This patient has good forearm movements of 90° of pronation and 80° of

supination and hand grip strength of the affected side of 75% of the opposite side. Patient has a MSTS score of 28, suggestive patient is having good functional upper extremity in terms of function, hand position, manual dexterity, emotional acceptance, and lifting abilities (Fig. 2).

This is another case that has undergone en bloc resection of the distal end of the radius and ulnar transposition. The post-operative period was uneventful. She was followed up and serial X-rays showed union of ulno-radial and ulno-carpal junction at the 4th month and 6th month, respectively. Her MSTS score was 26, suggesting she was able to do daily activities without much limitations. A year later, when she came in for follow-up, an X-ray was taken which showed a broken implant (Fig. 3). However, the patient was apparently normal and was able to perform daily activities without any pain or difficulties. The range of movements of the forearm was assessed for this patient which showed pronation of 90 degree and supination of 90 degree. This shows that almost the full range of pronation and supination was preserved in the forearm. Hand grip strength was assess showed 61% (20 kg) compared to the opposite side (34 kg). As a result of a broken implant, she can have jog dorsiflexion and palmar flexion at her wrist.

A limitation of our study is that it is a single institutional study, has no comparison group, and the sample size is limited.

Conclusion

This study confirms that wide excision and ulnar translocation are an effective surgical strategy for Campanacci Grade III GCT s of the distal radius. The procedure provides excellent oncological control with a low recurrence rate compared to intralesional surgery. Functionally, it allows for a painless, stable wrist while preserving a functional range of forearm rotation, resulting in high MSTS scores. Although grip strength is reduced, it remains sufficient for daily activities. The primary complication of non-union at the ulno-radial junction underscores the need for technical precision and robust internal fixation. For patients in resource-limited settings or those requiring biological durability, ulnar translocation remains a superior alternative to allografts, prosthetics, or distant autografts.

Clinical Message

In cases of aggressive distal radius giant cell tumors (Grade III), en bloc resection is the preferred oncological approach. Ipsilateral ulnar translocation is a reliable, biological reconstructive technique that preserves forearm rotation and achieves high functional outcomes. Surgeons should consider prophylactic bone grafting at the ulno-radial junction to prevent non-union and implant failure.

Declaration of patient consent: The authors certify that they have obtained all appropriate patient consent forms. In the form, the patient has given the consent for his/ her images and other clinical information to be reported in the journal. The patient understands that his/ her names and initials will not be published and due efforts will be made to conceal their identity, but anonymity cannot be guaranteed.

Conflict of interest: Nil **Source of support:** None

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