# Hexapod Circular Frame Fixation for Tibial Non-union: A Systematic Review of Clinical and Radiological Outcomes 

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#### Abstract

Introduction: Tibial non-unions present with complex deformities, bone loss, infection, leg length discrepancy (LLD), and other features which influence function. Circular frame-based treatment is popular with the hexapod system used increasingly. This systematic review aims to determine the clinical and radiological outcomes of hexapod fixation when used for tibial non-unions. Materials and methods: The review was performed in accordance with preferred reporting items for systematic reviews and meta-analyses (PRISMA) guidelines. The search strategy was applied to MEDLINE and Embase databases on 15 December 2021. Studies reporting either clinical or radiological outcomes following hexapod fixation on tibial non-unions were included. Primary outcomes were radiological union and patient-reported outcome measures (PROMs). Secondary outcomes included LLD, tibial alignment deformity (TAD), return to pre-injury activity and post-operative complications. Results: After the abstract and full-text screening, 9 studies were included; there were 283 hexapod frame fixations for tibial non-unions. Infection (46.6\%) and stiff hypertrophic non-union (39.2\%) accounted for most non-unions treated. The average age and mean follow-up were 42.2 years and 33.1 months, respectively. The average time to union was 8.7 months with a union rate of $84.8 \%$. A total of $90.3 \%$ of patients had TAD below $5^{\circ}$ in all planes, with an LLD $\leq 1.5 \mathrm{~cm}$ of the contralateral leg in $90.5 \%$. Bony and functional results were at least good in over $90 \%$ of patients when using the Association for the Study of the Method of llizarov (ASAMI) criteria. A total of 84\% of patients returned to pre-injury activities. There were complications as follows: a total of $34 \%$ developed pin-site infection, almost $9 \%$ experienced half-pin breakage and $14 \%$ developed an equinus ankle contracture. Conclusion: Hexapod frames for the treatment of tibial non-unions produce favourable functional outcomes. Complication rates are present and need to be discussed when this modality of treatment is proposed. Further comparative studies will allow for this option to be evaluated against that of the traditional llizarov frame and other methods of non-union surgery.


Keywords: Deformity Correction, Functional outcomes hexapod, Non-union, Union.
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## Introduction

Tibial non-union, following a fracture, can range between 2.5 and $17.5 \%$ owing to its anatomical location and soft-tissue coverage. ${ }^{1,2}$ Non-union can be classified as hypertrophic or atrophic, stiff or mobile, with or without bone defects and the presence or absence of infection. ${ }^{3,4}$ Such diversity underlines the range of different treatment strategies and why there currently is no standardised method for its management. ${ }^{5,6}$ However, a recent treatment algorithm for the use of circular frames in distraction, deformity correction, stabilisation and bone transport based on the type of tibial non-union has been proposed. ${ }^{7}$ Although the llizarov method has many advocates over the last three decades for hypertrophic non-union, ${ }^{8-11}$ and non-unions associated with bone defect and infection, ${ }^{12-16}$ there are limitations in its use. Despite allowing for simultaneous distraction and compression, it has significant learning curves with frequent modifications, ${ }^{12,13,17}$ need for multiple sequential corrections for angulation, translation and rotational deformities, ${ }^{9,11,18,19}$ and a protracted time in frame with concerns of pin-site infection. ${ }^{20,21}$ Such factors contribute to its increased costs. ${ }^{22,23}$ Hexapod frames are a modification of the llizarov-type fixators. ${ }^{24}$ Whilst applying the llizarov principles of distraction osteogenesis, ${ }^{25}$ they use specialised struts and computer programme to calculate the position of a virtual hinge to simultaneously correct the multiplanar deformities without altering
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the frame construct during treatment. ${ }^{18,26-28}$ Thus, compared to the llizarov method, they have a higher degree of precision for deformity correction and lower limb mechanical axis re-alignment, and a clear advantage in multidimensional deformity corrections. ${ }^{27,29}$ This is particularly useful for tibial non-unions frequently presenting with complex deformities, bone loss, infection and LLD, factors which can affect the union. ${ }^{30}$

In view of this, we performed a systematic review of the literature to investigate the clinical and radiological outcomes of hexapod frames on tibial non-unions.

## Materials and Methods

## Literature Search

A systematic review of the literature was conducted in accordance with the PRISMA guidelines, ${ }^{31}$ using the online databases MEDLINE and Embase. This was conducted from the inception of the databases to 15 December 2021. The full search strategy can be found in Appendix 1. No restriction was made on language with efforts made to obtain the translated-to-English versions of all included studies. Bibliographies of included studies were examined for missed and potentially relevant studies.

## Eligibility Criteria

All titles and abstracts returned by the search strategy were screened to identify studies reporting on clinical and radiological outcomes of hexapod frames on tibial non-unions. The main outcomes were as follows: (a) Clinical and radiological union and time to union; (b) TAD; (c) LLD; (d) function including a return to pre-injury work activity; and (e) post-surgical complications, namely, pin-site infection, component breakage, equinus contracture, LLD $>1.5 \mathrm{~cm}$ and regenerate site deformity. Exclusion criteria were paediatric population, non-human studies, case reports or expert opinions, foreign papers not translatable to English and those involving intra-articular regions.

## Study Selection and the Assessment of Quality

Two authors ( $K B$ and $S K$ ) independently reviewed the titles and abstracts, after which the relevant papers were reviewed in full by each author. Those that met the eligibility criteria were chosen with any discrepancies reviewed by a third author (AA). The same two authors independently assessed the quality of studies using the modified Coleman methodology score (MCMS) adjusted to account for the subject matter (Table 1). ${ }^{32}$ The MCMS is based on a scale ranging $0-100$; scores of $85-100$ are considered excellent, $70-84$ are considered good, 55-69 are considered fair, and scores below 55 are considered poor. ${ }^{32}$ Any discrepancy of more than 4 points between both reviewers was highlighted and resolved by the senior author (AA).

## Results

A total of 216 abstracts were identified from the initial search. Application of the eligibility criteria resulted in the inclusion of 9 studies. ${ }^{25,33-40}$ This is summarised in Flowchart 1. After full data extraction, we recognised there was insufficient data to undertake a meta-analysis. We, therefore, proceeded to do a qualitative synthesis of the data.

## Methodological Quality of Included Studies

Regarding MCMS of the 9 studies (mean score: 59.7), 6 achieved fair scores, ${ }^{33-36,38,39}$ and 3 poor scores. ${ }^{25,37,40}$ The overall quality of the studies was fair. Baseline characteristics are provided in Table 2.

In total there were 283 hexapod frame fixations for tibial non-unions. The averageageandmeanfollow-upswere 42.2 years and 33.1 months, respectively. Common methods of fracture stabilisation before limb salvage with hexapod fixation were monolateral external fixators (25.4\%), plate osteosynthesis (24.3\%) and intramedullary nailing (19.8\%). The average number of surgeries before hexapod fixation was 2.6. The Taylor spatial frame (TSF) (Smith and Nephew, Inc, Memphis, Tennessee) was the predominant hexapod used ( $77.4 \%$ ), with $9.2 \%$ of patients treated with the TrueLock-Hex
(TL-HEX) (Orthofix, Verona, Italy) and the remainder unknown. ${ }^{33}$ Stiff hypertrophic non-unions (39.2\%) and infected non-unions (46.6\%) accounted for most cases treated with the hexapod, with the former undergoing closed distraction or deformity correction or both, and the latter bone transport. Furthermore, the closed distraction was predominantly monofocal. ${ }^{35-38}$ Bifocal osteogenesis was generally performed for bone transport, ${ }^{25,33,34}$ with trifocal performed for larger defects. ${ }^{39}$ There were $40.7 \%$ of patients who were smokers at the time of hexapod fixation.

## Radiological Outcomes (Table 3)

The average time to union was 8.7 months. Subgroup analysis revealed those with infected non-unions united at 10.6 months, $25,33,34,40$ compared to 5.8 and 5.6 months in those without infection and with stiff hypertrophic non-unions, respectively. ${ }^{25,35,37,40}$ There was union in $84.8 \%$ of all cases after hexapod fixation, with $97.1 \%$ uniting after adjuvant stability was introduced. The remaining 6 patients either had an amputation (four cases) ${ }^{25,37,38}$ or withdrew from treatment, ${ }^{36}$ or were erroneously treated with closed distraction. ${ }^{35}$ Subgroup analysis revealed 6 studies that specifically gave union rates for infected non-unions. A reported $100 \%$ union was achieved in four studies. ${ }^{36,38-40}$ One study revealed infection to be an independent risk factor for non-union. ${ }^{25}$ The final study showed union in $55.2 \%$ of cases but did not record results after adjuvant stabilisation. ${ }^{33}$ Three studies reported on union rates for stiff hypertrophic non-union: $100 \%$ union was achieved in one, ${ }^{40}$ with a $98 \%$ union from $87 \%$ after adjuvant stability in the other two. ${ }^{35,37}$

Absence of malalignment was recorded in $90.3 \%$ of patients where a TAD of $<5^{\circ}$ in all planes and an LLD $\leq 1.5 \mathrm{~cm}$ to the contralateral leg in $90.5 \% .^{25,34-39}$ Only one study compared LLD before and after hexapod application, with over 1-cm improvement in deformity. ${ }^{25}$

## Clinical Outcomes (Table 4)

The ASAMI scores revealed bony and functional results to be at least good in 94.2 and $90 \%$ of patients, respectively. $25,34,38,39$ The 12-item short-form health survey (SF-12) in two studies revealed the patient's physical and mental scores to be within the norm of the US population mean score (50). ${ }^{37,40}$ One study used the short musculoskeletal functional assessment (sMFA) tool and found worse function after hexapod application compared to the standard population (27.1 vs $12.7, p<0.0001$ ). ${ }^{33}$ There were $84.2 \%$ of patients who were able to return to pre-injury activities. ${ }^{34,38,40}$ Subgroup analysis revealed smokers had slightly worse SF-12 physical scores $(47.89 \pm 14.13 \text { vs } 50.09 \pm 7.00)^{37}$ and sMFA scores ( $39 \pm 16$ vs $22 \pm$ $14, p=0.011) .{ }^{33}$

## Complications (Table 5)

Six studies reported on pin-site infection, ${ }^{34,35,37-40}$ although one presented its overall data to include llizarov frames. ${ }^{39}$ After exclusion, $34.3 \%$ of patients developed a pin-site infection following hexapod fixation for tibial non-unions. The same six studies reported half-pin breakage in $8.8 \%$ of all cases. Two studies commented on the development of equinus contractures in $14.3 \% .^{34,39}$ These patients had trifocal bone transport and underwent successful treatment with Achilles tendon lengthening with frame extension to the foot. There were $9.5 \%$ of patients who had LLD above 1.5 cm and were treated with a shoe lift. Three studies reported on regenerate site bending, with above

Table 1: Modified Coleman's criteria used for assessment of the quality of studies

| Criteria |  |  |
| :---: | :---: | :---: |
| Part A |  |  |
| Study size (total patients) | $>40$ | 15 |
|  | 25-40 | 10 |
|  | 11-24 | 5 |
|  | <10 | 0 |
| Mean follow-up (months) | $>24$ | 10 |
|  | 12-24 | 5 |
|  | <12, not stated or unclear | 0 |
| Type of study (methodology) | Randomised controlled trial | 12 |
|  | Prospective cohort study | 7 |
|  | Prospective/retrospective mixed | 3 |
|  | Retrospective cohort study | 0 |
| Diagnostic certainty (confirmed non-union) | In all | 5 |
|  | >80\% | 3 |
|  | <80\% | 0 |
| Part B |  |  |
| Outcome criteria (15) | Clearly defined outcome | 3 |
|  | Timing of outcome assessment clearly stated | 3 |
|  | PROMs used | 3 |
|  | Radiological assessment | 3 |
|  | Other clinical/functional outcomes measured (other than PROMs) | 3 |
| Procedures for assessing outcomes (6) | Clearly defined | 2 |
|  | Objective | 2 |
|  | Multiple/independent observers | 2 |
| Description of subject population (10) | Inclusion criteria reported and unbiased | 4 |
|  | Recruitment rate reported $>80 \%$ | 3 |
|  | Recruitment rate reported $<80 \%$ | 2 |
|  | Recruitment rate not reported | 0 |
|  | All eligible subjects accounted for in methodology | 3 |
| Surgical technique (6) | Method of bone transport | 2 |
|  | llizarov principles adequately described | 2 |
|  | Intraoperative techniques adequately described | 2 |
|  | Removal of frame justified | 2 |
| Post-operative rehabilitation (6) | Well described | 6 |
|  | Inadequately described | 3 |
|  | Protocol not reported | 0 |
| Complications recorded (8) | All with explanations | 8 |
|  | Selected complications recorded | 4 |
|  | Incomplete record | 2 |
|  | None | 0 |
| Adjuvant stability (5) | Time to adjuvant | 3 |
|  | Stabilisation method | 2 |
|  | Nil adjuvant | 3 |

$5^{\circ}$ occurring in $5.7 \%$ of cases. ${ }^{34,38,39}$ All were successfully treated with secondary correction.

## Discussion

Management of tibial non-unions can be challenging. The hexapod system incorporates the llizarov technique of translating controlled axial micromovements into a biomechanical environment conducive to bone healing and regenerate formation. ${ }^{26}$ In addition, they have
a much higher degree of precision for deformity correction. ${ }^{27,29}$ By adjusting the length of the six connecting struts, deformity correction can be achieved simultaneously with bone transport, and whilst restoring limb length discrepancy, eradicating infection, achieving union and soft-tissue coverage. This restores a functional limb and limits further complications. ${ }^{12,14}$

Our systematic review confirms the effective management of tibial non-unions with the hexapod frame. It reliably promotes

Flowchart 1: Preferred reporting items for systematic reviews and meta-analyses flowchart

union whilst accurately correcting concurrent deformities and limb-length discrepancies.

Patients with infected non-unions took 4.8 months longer to unite than those without, with Rozbruch et al. showing infection itself to be a risk factor for non-union. ${ }^{25}$ This can be explained as follows: first, bone transport using the hexapod system was undertaken in the majority of these cases due to significant bone defects following debridement and grafting. Bone transport is inherently more complicated than compression distraction, with respectively longer treatment times and further operative procedures necessary. Additionally, a time delay exists before bony contact and compression at the docking site, which adds to treatment time. ${ }^{13}$ A second reason is that, with any infected case, time to union is generally longer and more difficult. ${ }^{15}$ Third, there are potential confounding factors for which the duration of hexapod fixation may depend on, for example, the patient's immune status, comorbidities, and type and chronicity of the infection.

The restoration of LLD within 1.5 cm (90.5\%) and TAD $<5^{\circ}$ in all planes ( $90.3 \%$ ) underlines the simplicity of using a hexapod and easier control of bony re-positioning than when using a traditional llizarov frame. This ease of use is cited by other authors as advantage of the system. ${ }^{41,42}$ Five degrees of TAD in the sagittal (apex posterior or anterior angulation) or coronal (varus or valgus) was the value chosen in all included studies as acceptable for a normal mechanical axis. ${ }^{43}$ Any significant alterations may increase the joint reaction forces leading to potential progression in knee and ankle osteoarthritis. ${ }^{43}$ However, full correction of tibial alignment and LLD was not achieved in some studies. ${ }^{25,34,35,37}$

Although PROMs were generally favourable, Napora et al. showed worse mean sMFA scores than the uninjured reference population (27.1 vs $12.7, p<0.0001$ ). ${ }^{33}$ The context to consider is the residual dysfunction that may remain for most of these patients following such complex musculoskeletal injury. There were improved sMFA scores at 8 years ( 19.4 at 98.8 months vs 27.1 at 59 months), suggesting that over time these patients approach levels of that of the normal population. Interestingly, univariate analysis of smoking
in this study and that by Mahomed et al. ${ }^{37}$ showed worse sMFA and SF-12 scores at the final follow-up respectively. This may suggest that links to delayed healing, higher non-union rates, and altered biomechanical properties of bone with nicotine exposure. ${ }^{44}$ These findings are consistent with a previous study. ${ }^{45}$

Complication rates may be considered reasonable owing to the complex patient population, with pin-site infection rates similar to established series of frame management of tibial defects and deformity. ${ }^{13}$ All cases responded to treatment with oral antibiotics, except for three in which two required wire re-positioning ${ }^{34,38}$ and one required debridement, irrigation and hexapod re-application following deep infection. ${ }^{40}$ Equinus contractures at the ankle are common complications during tibial lengthening and bone transport, particularly in trifocal transport, as underlined in the two included studies. ${ }^{34,39}$ Despite successful treatment with Achilles tendon lengthening and frame extension to the foot, such complications can be prevented in the future through incorporating the foot in fixation for lengthening of more than $10 \%{ }^{46}$

## Limitations

A meta-analysis was not conducted owing to the heterogeneity of the methods, the limited quality of several studies (three studies were of poor quality) and small sample sizes. The latter is understandable due to the scarcity and complexity of the patient population. A further limitation includes most of the studies performed at centres specialising in limb reconstruction, thereby limiting its external validity. Therefore, the results reported may be difficult to replicate in less experienced trauma units. However, treatment of tibial non-unions requires experience and specialised knowledge to achieve satisfactory outcomes.

## Conclusion

This systematic review suggests hexapod frames are reliable for treating tibial non-unions with favourable outcomes. Future comparative studies should be undertaken to prove its efficacy over that of the llizarov frame.
Table 2: Baseline characteristics of included studies

| Study | Study design | Study participants | Gender |  | Mean age $\pm$ SD (range) | Initial fracture: <br> Closed vs open <br> (\%) | Initial surgery before hexapod |  | Hexapod |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  | Male | Female |  |  | Type of surgery | Average number of surgeries | TSF | TL-HEX | Other |
| Napora et al. ${ }^{33}$ | Retrospective cohort | 38 | 28 | 10 | $46.8 \pm 12.7$ | Not described | Not described | Not described | - | - | 38 |
| Sala et al. ${ }^{34}$ | Retrospective case | 12 | 8 | 4 | 44 (19-79) | 0 vs 12 (100) | PO: 3, IM nail 2, Ex-fix: 7 | 3 | 12 | - | - |
| Ferreira et al. ${ }^{35}$ | Prospective | 44 (46 frames) | 31 | 13 | 35 (18-68) | $\begin{aligned} & 10(21.7) \text { vs } 36 \\ & (78.3) \end{aligned}$ | Cast: 9, IM nail: 1 Unknown: 36 | Not described | 24 | 22 | - |
| Rozbruch et al. ${ }^{25}$ | Retrospective case | 38 | 30 | 8 | 43 (8-72) | 10 (26.3) vs 26 (68.4) 2 bone defects | PO: 9, IM nail:10, $\text { Ex-fix } 19$ | 4 | 38 | - | - |
| Arvesen et al. ${ }^{36}$ | Retrospective case | 34 (37 frames) | 26 | 11 | 49.7 (29-71) | 7 (18.9) vs 30 (81.9) | PO, IM nail and Ex-fix | Not described | 37 | - | - |
| Mahomed et al. ${ }^{37}$ | Retrospective case | 32 (33 frames) | 24 | 8 | 44 (18-80) | Not described | Cast: 7, PO: 7, IM nail: 10, Ex-fix: 8 | Not described | 29 | 4 | - |
| Khunda et al. ${ }^{38}$ | Retrospective case | 40 | 28 | 12 | 39.5 (9-69) | 16 (40) vs 24 (60) | Cast: 2, PO: 15, IM nail: 12, Ex-fix: 11 | 2 | 40 | - | - |
| Aboumira et al. ${ }^{39}$ | Retrospective cohort | 55 (30 with TSF) | 25 | 5 | $\begin{array}{r} 39 \pm 20.4 \\ (15-79) \end{array}$ | Not described | Not described | Not described | 30 | - | - |
| Molepo et al. ${ }^{40}$ | Retrospective case | 9 | 7 | 2 | $38 \pm 13$ | Not described | PO: 9 | 1 | 9 | - | - |


| Non-union |  |  |  |  |  | Method of distraction osteogenesis | Mean size of defect (cm) | Mean external fixator index (months/cm) | Mean follow-up (months) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Study | Type | Tibial site | Treatment | Risk factor: Smoking $[n,(\%)]$ | Risk factor: Diabetes [ $n,(\%)$ ] |  |  |  |  |
| Napora et al. ${ }^{33}$ | Infection | Not described | Bone transport | 11 (29) | 3 (8) | Bifocal | $5.1 \pm 1.6$ | $1.9 \pm 0.06$ | 59 |
| Sala et al. ${ }^{34}$ | Infection | Proximal: 2 <br> Middle: 4 <br> Distal: 6 | Bone transport | - | - | Bifocal: 6 <br> Trifocal: 6 | - | $\begin{aligned} & 2.0 \pm 0.9 \\ & \text { (Bi: } 2.63, \text { Tri } 1.31 \text { ) } \end{aligned}$ | $24 \pm 4.7$ |
| Ferreira et al. ${ }^{35}$ | Stiff hypertrophic | Not described | Closed distraction and deformity correction | 19 (43.2) | 1 (2.3) | Monofocal: 46 | - | - | $\begin{aligned} & 12 \\ & (6-40) \end{aligned}$ |
| Rozbruch et al. ${ }^{25}$ | Hypertrophic: 6 <br> Atrophic 18 <br> Normotrophic 14 <br> Infected: 19 | Proximal: 6 <br> Middle: 12 <br> Distal: 20 | Hypertrophic: Closed distraction Atrophic: Mechanical alignment, bone graft, compression Infected: Bone transport | 10 (26.3) | 4 (10.5) | Bifocal 17 <br> Trifocal: 3 <br> Unknown in rest | 6.5 (1-16) | - | $\begin{aligned} & 37 \\ & (15-63) \end{aligned}$ |
| Arvesen et al. ${ }^{36}$ | Hypertrophic: 24 Atrophic: 9 Normotrophic: 4 Infection 21 | Distal: 37 | Hypertrophic: Closed distraction <br> Atrophic: Mechanical alignment, bone graft, compression Infected: Bone transport | 20 (54.1) | 2 (5.4) | Predominantly monofocal Bifocal if degenerate deficient | - | - | 25.1 |
| Mahomed et al. ${ }^{37}$ | Stiff hypertrophic | Proximal: 6 <br> Middle: 6 <br> Distal: 21 | Closed distraction and deformity correction | 20 (62.5) | 10 (31.3) | Monofocal | - | - | Does not state |


| Khunda et al. ${ }^{38}$ | Infection: 15 <br> Unknown: 25 | Not described | Not adequately described | 12 (30) | - | Monofocal 25 Bifocal: 15 | - |  |  | 26 (3-70) |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Aboumira et al. ${ }^{39}$ | Infection: 20 <br> Unknown: 10 | Not described | Bone transport | - | - | Bifocal: 10 <br> Trifocal: 20 | $\begin{aligned} & 7.6 \pm 3.5 \\ & (3-15) \end{aligned}$ | $\begin{aligned} & 1.97 \pm 0.7 \\ & (1.1-3.4) \end{aligned}$ |  | $48 \pm 12.8$ |  |
| Molepo et al. ${ }^{40}$ | Infection: 7 Hypertrophic: 2 | Distal: 9 | Bone transport Closed distraction and deformity correction | - | - | Not described | - | - |  | $\begin{aligned} & 41.7 \pm \\ & 28.3 \end{aligned}$ |  |
|  | Outcomes |  |  | Rehabilitation protocol |  |  |  |  | CMS |  | Value |
| Study |  |  |  | A | $B$ | Total |  |
| Napora et al. ${ }^{33}$ | sMFA |  |  |  |  |  |  | Day 0: Weight-bearing as tolerated |  |  |  | 35 | 29 | 64 | Fair |
| Sala et al. ${ }^{34}$ | Mechanical axis deviation, LLD, Radiological union, ASAMI, Return to work |  |  | Day 0: Isometric quadriceps and knee ROM exercise Day 2: PWB Dynamisation 2 weeks before frame removal Week 4-6 post-frame removal: PWB |  |  |  | 20 | 43 | 63 | Fair |
| Ferreira et al. ${ }^{35}$ | TAD, LLD, Radiological union |  |  | Dynamise after consolidation FWB after this period |  |  |  | 32 | 37 | 69 | Fair |
| Rozbruch et al. ${ }^{25}$ | TAD, LLD, Radiological union, SF-36, AAOS, ASAMI, |  |  | Not described |  |  |  | 25 | 29 | 54 | Poor |
| Arvesen et al. ${ }^{36}$ | Deformity correction in 6 axes, Mechanical axis deviation, LLD, Radiological union |  |  | Day 0: weight-bearing as tolerated Dynamize after consolidation FWB 2-3/52 before frame removal |  |  |  | 25 | 33 | 58 | Fair |
| Mahomed et al. ${ }^{37}$ | TAD, Angulation, LLD, Radiological union, SF-12 form, |  |  | Day 0: Weight-bear as tolerated with early ROM exercises |  |  |  | 15 | 38 | 53 | Poor |
| Khunda et al. ${ }^{38}$ | ASAMI, Satisfaction score, Radiological union, Return to work |  |  | Not described |  |  |  | 25 | 38 | 63 | Fair |
| Aboumira et al. ${ }^{39}$ | Radiological union, LLD, ASAMI |  |  | Day 0: Quadriceps isometric and Knee ROM exercises <br> Day 2: PWB with crutches <br> Dynamize 3 weeks before frame removal <br> Week 4-6 post-frame removal: PWB Week 6 onwards post-frame: <br> FWB |  |  |  | 20 | 42 | 62 | Fair |
| Molepo et al. ${ }^{40}$ | SF-12, Foot function index, Radiological union, Return to work |  |  | Not described |  |  |  | 15 | 36 | 51 | Poor |

[^0]Table 3: Radiological outcome measures in hexapod fixation for tibial non-unions

| Study | Ilizarov principles |  |  | Mean time in frame/time to union (months $\pm$ SD) | Radiological |  |  |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  |  | Lower limb axis measurements | $T A D<5^{\circ}$ in all planes[n, (\%)] | Leg length alignment ( $\leq 1.5 \mathrm{~cm}$ of contralateral leg) (\%) | Union |  |
|  | Latency (days) | Distraction rate (mm/day) | Consolidation (days) |  |  |  | $\begin{aligned} & \text { mMPTA } 87^{\circ} \\ & (85-90) \\ & \hline \end{aligned}$ | $\begin{aligned} & \text { aPPTA81} \\ & (77-84) \\ & \hline \end{aligned}$ | $\begin{aligned} & m L D T A 89^{\circ} \\ & (86-92) \\ & \hline \end{aligned}$ | $\begin{aligned} & \text { aADTABO } \\ & (78-82) \\ & \hline \end{aligned}$ | Post-hexapod Fixation (\%) | Adjuvant stability (\%) |
| Napora et al. ${ }^{33}$ | - | $\begin{gathered} 0.75(\mathrm{NS}) \\ 0.5(\mathrm{~S}) \end{gathered}$ | - |  | $9.3 \pm 3.0$ | - | ${ }^{-}$ | $-$ | ${ }^{-}$ | - | - | 21 (55.2) | Unknown post-re-application of hexapod, IM nail or PO |
| Sala et al. ${ }^{34}$ | 12-14 | 0.5-1.0 | - | $13.9 \pm 3.3$ <br> (Bifocal: 15.2 <br> Trifocal: 12.6) | Pre: 90.2 <br> Final: 87 | Pre: 83.3 <br> Final: <br> 80.5 | Pre: 92 <br> Final: 88 | Pre: 80.5 <br> Final: 80.1 | 12 (100) | 10 (83.3) | 12 (100) | Not required |
| Ferreira et al. ${ }^{35}$ | 0 | 1.0 | - | $\begin{aligned} & 5.75 \\ & (2.75-12.25) \end{aligned}$ | - | - | - | - | 42 (91.3) | 46 (100) | 41 (89.1) | 45 (97.8) <br> post-re-application of hexapod in 4 that failed |
| Rozbruch et al. ${ }^{25}$ | - | - | - | $\begin{aligned} & 9.63 \\ & \text { (3.97-23.83) } \\ & \text { Infection > } \\ & \text { No infection } \\ & 11.47 \pm 5.73 \\ & \text { vs } 7.2 \pm 3.4 \text {, } \\ & p 0.02) \end{aligned}$ | - | - | - | - | 32 (84.2) | Pre-frame vs post-frame 3.1 cm (1-5.7) vs 1.8 cm (0-6.8) | 27 (71.1) <br> Non-union: <br> Infection > <br> No infection <br> [9/11 (81.1\%) <br> vs $2 / 11$ <br> (18.2\%), <br> $p=0.03$ ] | 36 (94.7) <br> post-TSF <br> re-application and ABG: 4, <br> IM nail: 3, PO: 2 and amputation: 2 |
| Arvesen et al. ${ }^{36}$ | - | 0.5-1.0 | - | $6.01 \pm 3.56$ | 36 (97.3\%) corrected within $5^{\circ}$ | - | 25 (67.6\%) corrected within $5^{\circ}$ | - | 37 (100) | 36 (97.3) | 32 (86.4) | $\begin{aligned} & 35(94 \%) \\ & \text { post-TSF } \\ & \text { re-application: } 3 \text {, } \\ & \text { TTC fusion: } 1 \\ & \text { Treatment refusal: } 1 \end{aligned}$ |
| Mahomed et al. ${ }^{37}$ | 10 | 1.0 | - | 5.27 | - | - | - | - | 24 (72.7) | 24 (72.7) | 29 (87.9) | 32 (97) post-ABG: 3 Amputation: 1 |
| Khunda et al. ${ }^{38}$ | - | ${ }^{-}$ | - | 10.5 (3-38.5) | - | - | - | - | - <br>  <br>  <br> 1000$)$ | ${ }^{-}$ | 39 (97.5) | Not required Amputation for pain: 1 |
| Aboumira et al. ${ }^{39}$ | 12-14 | 0.5-1.5 | 56 | $\begin{aligned} & 13.9 \pm 4.8 \\ & (5.6-25.7) \end{aligned}$ | - | - | - | - | 30 (100) | 27/30 (90) | 30 (100) | Not required |
| Molepo et al. ${ }^{40}$ | - | - | - | $7.76 \pm 4.9$ <br> Infection> No infection $9.14 \pm 5.16$ vs $4.56 \pm 2.37$ | - | - | - | - | - | - | 9 (100) | Not required |

[^1]Table 4: The PROMs in hexapod fixation for tibial non-unionss

Table 5: Post-surgical complications following hexapod circular frame fixation for tibial non-union

|  | Post-surgical complications following hexapod fixation |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Study | Pin-site infection (\%) | Half-pin breakage (\%) | Equinus ankle contracture (\%) | $L L D>1.5 \mathrm{~cm}$ (\%) | Regenerate site bending (\%) | Others (\%) |
| Napora et al. ${ }^{33}$ | - | - | - | - | - | - |
| Sala et al. ${ }^{34}$ | 10 (83.3) Treatment: Local care 5 , oral antibiotics 4, Wire re-tension 1 | 4 (33.3) | 3 (25), All trifocal Treatment: AT lengthening with frame extension to foot | 2 (16.7) Treatment: Shoe lift | $<5^{\circ}: 3(25)$ <br> Nil treatment | Peroneal pseudoaneurysm:1 (8.3) <br> Treatment: Embolisation |
| Ferreira et al. ${ }^{35}$ | 9 (19.5) Treatment: Local care and oral antibiotics | 1 (2.2) Treatment: Replacement | - | 0 (0) | - | - |
| Rozbruch et al. ${ }^{25}$ | - | - | - | - | - | - |
| Arvesen et al. ${ }^{36}$ | - | - | - | 1 (2.7) Treatment: Shoe lift | - | - |
| Mahomed et al. ${ }^{37}$ | 5 (15.2) Treatment: Local care and oral antibiotics | 2 (6.1) Treatment: Replacement | - | 9 (27.2) Treatment: Shoe lift | - | - |
| Khunda et al. ${ }^{38}$ | $23 \text { (57.5) }$ <br> Treatment: oral antibiotics 22, wire re-tension 1 | 1 (2.5) Treatment: Replacement | - | - | $>5^{\circ}: 2$ (5) Treatment: Secondary correction | Knee flexion deformity: 1 (2.5) Treatment: Frame extension above knee |
| Aboumira et al. ${ }^{39}$ | 31/55 (56.3) Value for TSF in isolation unknown | 5 (16.7) Treatment: Removal | 3 (10), All trifocal Treatment: AT lengthening with frame extension to foot | 3 (10) Treatment: Shoe lift | $\begin{aligned} & <5^{\circ}: 3(10)>5^{\circ}: 2(6.7) \\ & \text { Treatment:TSF } \\ & \text { re-application: } 1, \text { PO: } 1 \end{aligned}$ | - |
| Molepo et al. ${ }^{40}$ | 1/9 (11.1) <br> Treatment: aggressive debridement, irrigation and TSF re-application | 2 (22.2) Treatment: Replacement | - | - | - | - |

## References

1. Phieffer LS, Goulet JA. Delayed unions of the tibia. J Bone Joint Surg Am 2006;88(1):206-216. DOI: 10.2106/00004623-200601000-00026.
2. Tzioupis C, Giannoudis PV. Prevalence of long-bone non-unions. Injury 2007;38(Suppl. 2):S3-S9. DOI: 10.1016/s0020-1383(07)80003-9.
3. Weber B, Cech O. Pseudoarthrosis: Pathology, Biomechanics, Therapy, Results. Bern, Switzerland: Hans Huber Medical Publisher, 1976.
4. Wu CC, Chen WJ. A revised protocol for more clearly classifying a nonunion. J Orthop Surg 2000;8(1):45-52. DOI: 10.1177/ 230949900000800109.
5. Akhtar A, Shami A, Sarfraz M. Functional outcome of tibial nonunion treatment by Ilizarov Fixator. Ann Pak Inst Med Sci 2012;8(3):188-191. Corpus ID: 74404660.
6. Jones CB, Mayo KA. Nonunion treatment: Iliac crest bone graft techniques. J Orthop Trauma 2005;19(Suppl. 10):S11-S13. DOI: 10.1097/00005131-200511101-00004.
7. Ferreira N, Marais LC. Management of tibial non-unions according to a novel treatment algorithm. Injury 2015;46(12):2422-2427. DOI: 10.1016/j.injury.2015.09.040.
8. Catagni MA, Guerreschi F, Holman JA, et al. Distraction osteogenesis in the treatment of stiff hypertrophic nonunions using the Ilizarov apparatus. Clin Orthop Relat Res 1994;301:159-163. PMID: 8156667.
9. Ilizarov GA. Clinical application of the tension-stress effect for limb lengthening. Clin Orthop Relat Res 1990;250:8-26. PMID: 2403497.
10. Kocaoglu M, Eralp L, Sen C, et al. Management of stiff hypertrophic nonunions by distraction osteogenesis: A report of 26 cases. J Orthop Traum 2003;17(8):543-548. DOI: 10.1097/00005131-200309000-00001.
11. Saleh M, Royston S. Management of nonunion of fractures by distraction with correction of angulation and shortening. J Bone Joint Surg 1996;78(1):105-109. PMID: 8898138.
12. Cattaneo R, Catagni M, Johnston EE. The treatment of infected nonunions and segmental defects of the tibia by the methods of the Ilizarov. Clin Orthop Relat Res 1992;280:143-152. PMID: 1611734.
13. Paley D, Maar DC. Ilizarov bone transport treatment for tibial defects. J Orthop Trauma 2000;14:76-85. DOI: 10.1097/00005131-20000200000002.
14. Paley D, Catagni MA, Argnani F, et al. Ilizarov treatment of tibial nonunions with bone loss. Clin Orthop Relat Res 1989;241:146-165. PMID: 2924458.
15. Dendrinos GK, Kontos S, Lyritsis E. Use of Ilizarov technique for treatment of non-union of the tibia associated with infection. J Bone Joint Surg 1995;77(6):835-846. DOI: 10.2106/00004623-199506000-00004.
16. Marsh DR, Shah S, Elliott J, et al. The Ilizarov method in nonunion, malunion and infection of fractures. J Bone Joint Surg 1997;79(2): 273-279. DOI: 10.1302/0301-620x.79b2.6636.
17. Elbatray Y, Fayed M. Deformity correction with an external fixator: Easy of use and accuracy? Orthopaedics 2009;32:82. PMID: 19301808.
18. Feldman DS, Shin SS, Madan S, et al. Correction of tibial malunion and nonunion with six-axis analysis deformity correction using the Taylor spatial frame. J Orthop Trauma 2003;17(8):549-554. DOI: 10.1097/00005131-200309000-00002.
19. Shtarker H, Volpin G, Stolero J, et al. Correction of combined angular and rotational deformities by the llizarov method. Clin Orthop Relat Res 2002;402:184-195. DOI: 10.1097/00003086-20020900000017.
20. Garcia-Cimbrelo E, Marti-Gonzalez JC. Circular external fixation in tibial non-unions. Clin Orthop Relat Res 2004;419:65-70. PMID: 15021133.
21. Maini L, Chadha M, Vishwanath J, et al. The Ilizarov method in infected nonunion of fractures. Injury 2000:31(7):509-517. DOI: 10.1016/s0020-1383(00)00036-x.
22. Patil S, Montgomery R. Management of complex tibial and femoral nonunion using the llizarov technique, and its cost implications. J Bone Joint Surg Br 2006;88(7):928-932. DOI: 10.1302/0301620X.88B7.17639.
23. Williams MO. Long-term cost comparison of major limb salvage using the llizarov method versus amputation. Clin Orthop Relat Res 1994;301:156-158. PMID: 8156666.
24. Fadel M, Hosny G. The Taylor spatial frame for deformity correction in the lower limbs. Int Orthop 2005;29(2):125-129. DOI: 10.1007/s00264-004-0611-9.
25. Rozbruch SR, Pugsley JS, Fragomen T. Repair of tibial nonunions and bone defects with the Taylor spatial frame. J Orthop Trauma 2008:22(2):88-95. DOI: 10.1097/BOT.0b013e318162ab49.
26. Rozbruch SR, Fragomen AT, Ilizarov S. Correction of tibial deformity with use of the Ilizarov-Taylor spatial frame. J Bone Joint Surg 2006;88(Suppl. 4):156-174. DOI: 10.2106/JBJS.F. 00745.
27. Manner HM, Huebl M, Radler C, et al. Accuracy of complex lowerlimb deformity correction with external fixation: A comparison of the Taylor spatial frame with the llizarov ring fixator. J Child Ortho 2007;1:55-61. DOI: 10.1007/s11832-006-0005-1.
28. Docquier PL, Rodriguez D, Mousny M. Three-dimensional correction of complex leg deformities using a software assisted external fixator. Acta Orthop Belg 2008;74(6):816-822. PMID: 19205330.
29. Rozbruch SR, Segal K, Ilizarov S, et al. Does the Taylor spatial frame accurately correct tibial deformities? Clin Orthop Relat Res 2010;46(5)8:1352-1361. DOI: 10.1007/s11999-009-1161-7.
30. Schartsman V, Choi SH, Schwartsman R. Tibial nonunions. Treatment tactics with the Ilizarov method. Orthop Clin North Am 1990;21(4):639-653. PMID: 2216399.
31. Liberati A, Altman DG, Tetzlaff J, et al. The PRISMA statement for reporting systematic reviews and meta-analyses of studies that evaluate health care interventions: explanation and elaboration. J Clin Epidemiol 2009;6(7):e10000100. DOI: 10.1371/journal. pmed. 1000100.
32. Coleman BD, Khan KM, Maffuli N, et al. Studies of surgical outcome after patellar tendinopathy: Clinical significant of methodological deficiencies and guidelines for future studies. Victoria Institute of Sport Tendon Study Group. Scan J Med Sci Sports 2000;10(1):2-11. DOI: 10.1034/j.1600-0838.2000.010001002.x.
33. Napora JK, Weinberg DS, Eagle BA, et al. Hexapod stacked transport for tibial infected nonunions with bone loss: Long-term functional uutcomes. J Orthop Trauma 2018;32(1):e12-e18. DOI: 10.1097/ BOT. 0000000000001005 .
34. Sala F, Thabet AM, Castelli F, et al. Bone transport for postinfectious segmental tibial bone defects with a combined Ilizarov/Taylor spatial frame technique. J Orthop Trauma 2011;25(3):162-168. DOI: 10.1097/ BOT.Ob013e3181e5e160.
35. Ferreira N, Marais LC, Aldous C. Hexapod external fixator closed distraction in the management of stiff hypertrophic tibial nonunions. Bone Joint J 2015;97-B(10):1417-1422. DOI: 10.1302/0301620X.97B10.35504.
36. Arvesen JE, Watson JT, Israel H. Effectiveness of treatment for distal tibial nonunions with associated complex deformities using a hexapod external fixator: J Orthop Trauma 2017;31(2):e43-e48. DOI: 10.1097/BOT. 0000000000000726.
37. Mahomed N, O'Farrell P, Barnard AC, et al. Monofocal distraction treatment of stiff aseptic tibial nonunions with hexapod circular external fixation. J Limb Lengthen Reconstr 2017;3(2):101-106. DOI: 10.4103/jllr.jllr_31_16.
38. Khunda A, AI-Maiyah M, Earldey WGP, et al. The management of tibial fracture non-union using the Taylor spatial frame. J Orthop 2016;13(4):360-363. DOI: 10.1016/j.jor.2016.07.002.
39. Aboumira IEA, Sala F, Elbatrawy Y, et al. Distraction osteogenesis for tibial nonunion with bone loss using combined Iliarov and Taylor spatial frames versus a conventional circular frame. Strategies Trauma Limb Reconstr 2016;11(3):153-159. DOI: 10.1007/s11751-016-0264-4.
40. Molepo M, Barnard AC, Birkholtz F, et al. Functional outcomes of the failed plate fixation in distal tibial fractures salvaged by hexapod external fixator. Eur J Orthop Surg Traumatol 2018;28(8):1617-1624. DOI: 10.1007/s00590-018-2231-x.
41. Perren SM. Evolution of the internal fixation of long bone fractures. The scientific basis of biological internal fixation: Choosing a
new balance between stability and biology. J Bone Joint Surg 2002;84(8):1093-1110. DOI: 10.1302/0301-620x.84b8.13752.
42. Feldman DS, Madan SS, Ruchelsman DE, et al. Accuracy of correction of tibia vara: acute vs gradual correction. J Pediatr Orthop 2006;26(6):794-798. DOI: 10.1097/01.bpo.0000242375.64854.3d.
43. Graehl PM, Hersh MR, Heckman JD. Supramalleolar osteotomy for the treatment of symptomatic tibial malunion. J Orthop Trauma 1987;1(4):281-292. DOI: 10.1097/00005131-198701040-00003.
44. Hernigou J, Schuind F. Smoking as a predictor of negative outcome in diaphyseal fracture healing. Int Orthop 2013;37:883-887. DOI: 10.1007/s00264-013-1809-5.
45. Christiano AV, Pean CA, Konda SR, et al. Predictors of patient reported pain after lower extremity non-union surgery: The nicotine effect. Iowa Orthop J 2016;36:53-58. PMCID: PMC4910799.
46. Eldridge JC. Problems with substantial limb lengthening. Orthop Clin North AM 1991;22(4):625-631. PMID: 1945340.

## Appendix 1

| 1 | exp fracture fixation/ | 158,523 | 13 | hexapod.mp. | 1,065 | 25 | 24 or 8 | 34,000 |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2 | fractures, bone/ | 97,307 | 14 | "circular external fixator*". mp. | 534 | 26 | fracture non-union/ or fracture healing/ | 55,370 |
| 3 | 1 or 2 | 236,711 | 15 | Taylor spatial frame.mp. | 549 | 27 | 10 or 26 | 196,372 |
| 4 | tibia/ | 78,839 | 16 | TL-HEX.mp. | 18 | 28 | 25 and 27 and 17 | 201 |
| 5 | 3 and 4 | 6,330 | 17 | or/13-16 | 2,013 | 29 | 9 and 12 and 17 | 163 |
| 6 | tibia\$.ti. | 58,986 | 18 | tibia fracture/or tibia shaft fracture/ | 15,412 | 30 | 28 or 29 | 216 |
| 7 | fracture\$.tw. | 60,8181 | 19 | exp fracture treatment/or fracture healing/ | 15,1789 |  | OVID MEDLINE | 72 |
| 8 | 6 and 7 | 20,972 | 20 | fracture/ | 89,827 |  | OVID Embase | 142 |
| 9 | 5 or 8 | 25,232 | 21 | 19 or 20 | 22,8747 |  | Cochrane | 2 |
| 10 | [non union or non-union or nonunion or un-united or ununited or delayed union or union or (fractur* adj2 healing)].tw. | 169,400 | 22 | tibia/or tibia shaft/ | 79586 | 31 | remove duplicates from 30 | 153 |
| 11 | fractures, ununited/or fracture healing/ | 56,509 | 23 | 21 and 22 | 6,133 |  |  |  |
| 12 | 10 or 11 | 196,122 | 24 | 18 or 23 | 20,796 |  |  |  |


[^0]:    AAOS, American Academy of Orthopaedic Surgeons; ASAMI, Association for the Study of the Method of Ilizarov Criteria; Ex-fix, external fixator; FWB, full weight bearing; IM, intra-medullary; PO, plate
    osteosynthesis; PWB, partial weight; ROM, range of motion; SF, short form health survey; TL-HEX, TrueLock hexapod; TSF, Taylor spatial frame; Trii, trifocal, LLD, leg length discrepancy; SMFA, short musculoskeletal functional assessment

[^1]:    ABG, autologous bone graft; IM, intra-medullary; PO, plate osteosynthesis; TSF, Taylor spatial frame; TTC, tibiotalocalcaneal fusion; aPPTA, anatomic posterior proximal tibial angle; NS, non-smoker S, smoker; mLDTA, mechanical lateral distal tibial angle, aADTA, anatomic anterior distal tibial angle; mMPTA, mechanical medial proximal tibial angle

