Early results of starting a fracture liaison service for fragility hip fractures in a regional South African hospital

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Background. Fragility hip fracture is a rising pandemic, with the burden currently shifting to developing countries. A fracture liaison service (FLS) was implemented in September 2021 in a regional South African (SA) hospital.

Objective. To describe two groups of patients with surgically treated fragility hip fractures - those before and after implementation of the FLS - in terms of early mortality, refracture and readmission rates.

Methods. Patients aged >50 years who sustained hip fractures between January 2020 and June 2022 were considered; those with high-energy injuries, pathological and periprosthetic fractures were excluded. Associations between treatment group and mortality, readmission and refracture rates within the first year after surgery were investigated. Subgroup analyses were also conducted for specific risk factors.

Results. A total of 299 patients (mean (standard deviation) age of 74.5 (11.0) years, 68.2% female) were included, of whom 32.8% (n=98) received bisphosphonates. The overall 30-day, 90-day and 1-year mortality rates were 9.4% (n=28), 17.4% (n=52) and 29.8% (n=89), respectively, while 24.1% (n=72) of patients had unplanned readmissions to hospital, and 2.7% (n=8) refractured within the first year after surgery. No statistical difference in 30-day, 90-day or 1-year mortality rate, refracture incidence or readmission incidence was observed between patient groups. Time to surgery was shorter after the implementation of the FLS (median (interquartile range (IQR)) 30.8 (22.3 - 48.9) hours v. before (median (IQR) 42.5 (23.1 - 70.8) hours, *p*=0.039).

Conclusion. An FLS was successfully implemented in a regional SA hospital. Early mortality, refractures and readmission rates were low or comparable with the existing literature, and not affected by treatment within the FLS nor by bisphosphonate initiation. Treatment within the FLS, however, resulted in a decrease in time to surgery.

Keywords: fragility hip fracture, osteoporosis, fracture liaison service, bisphosphonates

S Afr Med J 2025;115(9):e2998. https://doi.org/10.7196/SAMJ.2025.v115i9.2998

Fragility hip fracture (FFH) is the most devastating consequence of osteoporosis, due to the resultant morbidity, mortality and economic impact.[1,2] While it is truly a global pandemic, the burden is now shifting to developing countries, mostly in Asia and sub-Saharan Africa (SSA). This is primarily due to ageing populations and changes in secular trends. [3-6] Although important advances have been made in SSA since the turn of the millennium pertaining to research and preventive strategies, significant challenges remain within the context of a quadruple disease burden.[6-8]

Fracture liaison services (FLSs), created in 2003 by McLellan et al.,[9] have gained popularity and have been endorsed by the International Osteoporosis Foundation's 'Capture the fracture' campaign. [10] They have proven efficacy in the comprehensive management of FFHs, especially in combination with anti-osteoporotic medications such as bisphosphonates. [11-14] Bisphosphonate treatment after FFH offers a good safety profile and leads to a reduction in refractures and allcause mortality. $^{[12,15-21]}$

An FLS was implemented in a regional South African (SA) hospital serving a predominantly rural community. This study aimed to describe the patient population before and after the implementation of an FLS in terms of three main outcomes: mortality, refracture and unplanned readmission within the first year after surgery. Secondary objectives included describing differences between patients treated with bisphosphonates v. those treated without bisphosphonates, and evaluating potential risk factors for mortality at 1 year following surgery.

Methods Study design

A retrospective cohort study was conducted that considered all patients with FFH treated surgically in a regional hospital in the Western Cape Province, SA, between January 2020 and June 2022. This period includes 20 months prior to and 10 months following FLS introduction.

Study setting and participants

This single-centre study was conducted in a regional hospital responsible for serving public sector patients in three districts in the Western Cape. The hospital's orthopaedic department manages orthopaedic trauma, emergencies and general elective orthopaedic conditions, including arthroplasty, but does not have any subspecialty

In September 2021, an FLS was implemented for patients sustaining osteoporotic fractures. This FLS is based on four pillars: (i) identification of patients at risk; (ii) evaluation for secondary causes of osteoporosis; (iii) initiation of intravenous bisphosphonates, vitamin D and calcium supplementation; and (iv) ensuring co-management and follow-up with endocrine and medical units for continuation of care. An FFH is diagnostic of osteoporosis and is an indication to start antiresorptive medication, [22] and therefore all patients with FFH qualified for osteoporosis treatment as part of the FLS. The role of the orthopaedic department was to treat the

presenting injury on its merit, identify and diagnose osteoporosis, perform initial biochemistry and blood investigations and initiate medical treatment. The patients were then referred to the regional hospital's medical outpatients' department and an endocrinology unit at a tertiary academic institution for the follow-up management of osteoporosis.

All patients undergoing hip fracture surgery from January 2020 until June 2022 were evaluated for inclusion. Patients <50 years of age, with pathological or periprosthetic fractures, chronic fractures, or high-energy mechanisms, or patients with incomplete data were excluded from the study. Demographic and clinical data were obtained from provincial electronic databases, patient records and mortality rates confirmed by the Department of Home Affairs.

The patient cohort was divided into two groups (pre FLS era and FLS era), depending on whether they were treated before (January 2020 - August 2021) or after (September 2021 - June 2022) implementation of the FLS. They were also divided into bisphosphonate and non-bisphosphonate groups based on whether or not they were initiated on bisphosphonates. All surgeries were performed by a senior specialist, or a trainee (registrar or medical officer) under specialist supervision. No procedures were performed after hours or without specialist oversight.

The primary outcome measures were 30-day, 90-day and 1-year mortality rates, and secondary outcome measures were 30-day and 1-year unplanned readmission rates, as well as 1-year refracture rates post surgery. An unplanned readmission was defined as any unforeseen readmission to the hospital, and included complications such as fracture-related infections, prosthetic dislocations and any and all other medical and surgical reasons for admission. Elective procedures and planned readmissions were excluded from this definition.

Statistical analysis

Data were analysed using Statistica version 14 (TIBCO, USA). Continuous data were evaluated for normality, and reported as mean (standard deviation (SD)) or median (interquartile range (IQR)), depending on distribution. Categorical data were summarised as frequencies and counts. A comparative analysis on demographic and baseline clinical variables was performed between patient groups (pre-FLS era v. FLS era; bisphosphonates v. non-bisphosphonates) to evaluate the presence of potential confounding variables using an independent t-test for continuous variables, or χ^2 or Fisher's exact tests for categorical variables. Thereafter, both groups were evaluated against the primary outcome measures using a χ^2 or Fisher's exact test. The α -level was set at 0.05.

Subgroup analyses were performed including fracture type (intracapsular v. extracapsular hip fractures), age groups (50 - 65, 65 - 80, >80 years), different American Society of Anesthesiologists (ASA) classes, and genders using χ^2 or Fisher's exact tests.

Ethical clearance

A waiver of informed consent was granted by the Health Research Ethics Committee of Stellenbosch Univerity (ref. no. N22/11/142) due to the minimal risk involved and the retrospective nature of the study.

Results

General characteristics

During the 2.5-year study period, 360 patients aged >50 years received hip fracture surgery, and were considered for inclusion. Of these, 61 were excluded, due to having high-energy fractures (n=29), pathological fractures (n=13), periprosthetic fractures (n=3), chronic fractures (n=4), or incomplete data (n=12). A total of 299 patients were therefore included in the study.

Baseline variables were similarly distributed between all study groups (Table 1). The majority (73.2%, n=219) of patients were referred from district hospitals and clinics; only about a quarter of patients (26.8%, n=80) were inhabitants of the town, referred via the hospital's emergency centre. The mean (SD) age was 74.5 (11.0) years (95% CI 73.2 - 75.7), with 68.2% female (n=204) and 31.8% male (n=95). The mean age of females was 75.8 (11.4) years, and the mean age of males was 71.6 (9.3) years. A total of 51.2% (n=153) of fractures were intracapsular (neck-of-femur fractures), while 48.8% (n=146) were extracapsular (pertrochanteric femur fractures). The surgeries performed included 145 (48.5%) hip replacements (arthroplasty) and 154 (51.5%) internal fixations. The median (IQR) time from diagnosis to surgery was 39.0 (22.5 - 67.4) hours, ranging between 4 and 330 hours. More than half (52.5%, n=157) of patients had severe systemic disease according to the ASA. During the study period, a total of 98 patients (32.8%) were initiated on bisphosphonate therapy. These all occurred after the implementation of the FLS, representing an increase from 0% to 83.8%. Patients in the two cohorts were similar with regard to known risk factors, including age and comorbidity status.

Time to surgery was shorter after the implementation of the FLS (median (IQR) 30.8 (22.3 - 48.9) hours) v. before (42.5 (23.1 - 70.8) hours, p=0.039). Consequently, time to surgery was also shorter in the bisphosphonate group (median (IQR) 28.3 (22.0 - 47.8) hours) than the non-bisphosphonate group (median (IQR) 42.4 (23.1 - 69.9) hours, p=0.031) (Table 1).

Clinical characteristics

The overall 30-day, 90-day and 1-year all-cause mortality rates were 9.4% (*n*=28), 17.4% (*n*=52) and 29.8% (*n*=89), respectively. Secondary fractures occurred in 2.7% of patients (n=8) within the first year after surgery. The 30-day and 1-year unplanned readmissions to hospital were 7.0% (n=21) and 24.1% (n=72), respectively. Neither the initiation of bisphosphonates nor treatment within the FLS context had a statistically significant effect on mortality, readmissions or refractures in this study population during the first year post surgery (Table 2).

Mortality at 1 year was similar (p=0.222) in males (34.7%) and females (27.5%), and between neck of femur fractures v. pertrochanteric fractures (32.0% v. 27.4%, p=0.448). Mortality rates were lower in patients aged <65 years than those ≥65 years (10.8% v. 35.0%, p<0.001), and in those with ASA classification 1 or 2 v. \geq 3 (16.9% v. 41.4%, *p*<0.001) (Table 3).

Subanalyses were performed for different fracture types (neck of femur v. pertrochanteric), age groups, ASA categories and gender. In the extracapsular (pertrochanteric) femur fracture group, a lower allcause mortality rate was observed during the FLS era at 90 days (8.2% v. 23.5%, p=0.024) and 1 year (18.0% v. 34.1%, p=0.039) (Table 4). For patients aged 65 - 80 years, lower all-cause mortality was observed at 90 days in the group treated with bisphosphonates compared with no bisphosphonates (6.7% v. 20.2%, p=0.048) (appendix Table S1; https://coding.samedical.org/file/2369). No further statistically significant associations were observed in the sub-analyses (appendix Tables S1 - S3).

Discussion

Standards for the management of FFH are well established internationally.[23-25] There are limited management guidelines originating from Africa.[1,26,27] These guidelines and treatment goals include surgery within 48 hours of admission, specialist surgical supervision, orthopaedic-geriatric comanagement and the evaluation for and treatment of osteoporosis. The 'quadruple disease burden' faced

| | | Group | | | Group | | |
|--------------------------------------|-----------------------|------------------------|----------------------------|-----------------|-----------------------|-----------------------|-----------------|
| Variable, mean (SD)* | All | Bisphosphonates (n=98) | No bisphosphonates (n=201) | <i>p</i> -value | Pre-FLS era (n=182) | FLS era (n=117) | <i>p</i> -value |
| Age, years | 74.5 (11.0) | 75.2 (10.2) | 74.1 (11.3) | 0.429 | 74.0 (11.3) | 75.2 (10.4) | 0.373 |
| Gender | | | | | | | |
| Female | 68.2 (204) | 74.5 (73) | 65.2 (131) | 0.114 | 65.4 (119) | 72.65 (85) | 0.205 |
| Male | 31.8 (95) | 25.5 (25) | 34.8 (70) | | 34.6 (63) | 27.35 (32) | |
| Charlson comorbidity index | 3.9 (1.7) | 3.9 (1.7) | 4.0 (1.8) | 0.920 | 4.0 (1.8) | 3.9 (1.6) | 0.820 |
| ASA category, mean (n) | | | | | | | |
| 1 | 11.7 (35) | 12.2 (12) | 11.4 (23) | 0.775 | 11.0 (20) | 12.8 (15) | 0.730 |
| 2 | 35.8 (107) | 38.8 (38) | 34.3 (69) | | 34.1 (62) | 38.5 (45) | |
| 3 | 48.2 (144) | 45.9 (45) | 49.3 (99) | | 50.0 (91) | 45.3 (53) | |
| 4 | 4.3 (13) | 3.1 (3) | 5.0 (10) | | 4.9 (9) | 3.4 (4) | |
| Vitamin D (nmol/L) | 46.5 (19.1) | 47.0 (19.4) | 43.4 (17.5) | 0.492 | 54.0 (-) | 46.4 (19.2) | 0.693 |
| Corrected calcium (mmol/L) | 2.3 (0.1) | 2.3 (0.1) | 2.3 (0.1) | 0.624 | 2.3 (0.1) | 2.3 (0.1) | 0.538 |
| Fracture type, mean (n) | | | | | | | |
| NOF | 51.2 (153) | 46.9 (46) | 53.2 (107) | 0.326 | 53.3 (97) | 47.9 (56) | 0.407 |
| Pertrochanteric femur fracture | 48.8 (146) | 53.1 (52) | 46.8 (94) | | 46.7 (85) | 52.1 (61) | |
| Time to surgery, hours, median (IQR) | 39.0 (22.5 - 67.4) | 28.3 (22.0 - 47.8) | 42.4 (23.1 - 69.9) | 0.031 | 42.5 (23.1 - 70.8) | 30.8 (22.3 - 48.9) | 0.039 |
| Surgery type, mean (n) | | | | | | | |
| Arthroplasty | 48.5 (145) | 44.9 (44) | 50.25 (101) | 0.392 | 51.1 (93) | 44.4 (52) | 0.287 |
| Internal fixation | 51.5 (154) | 55.1 (54) | 49.75 (100) | | 48.9 (89) | 55.6 (65) | |
| Bisphosphonate initiation | 32.8 (98) | 100.0 (98) | 0.0(0) | - | 0.0(0) | 83.8 (98) | - |

| | | | _ | Group | | | |
|--------------------------------|-----------|------------------------|----------------------------|-----------------|---------------------|--------------------|-----------------|
| Clinical characteristic, n (%) | All | Bisphosphonates (n=98) | No bisphosphonates (n=201) | <i>p</i> -value | Pre-FLS era (n=182) | FLS era (n=117) | <i>p</i> -value |
| 30-day mortality | 28 (9.4) | 9 (9.2) | 19 (9.5) | >0.999 | 18 (9.9) | 10 (8.6) | 0.839 |
| 90-day mortality | 52 (17.4) | 15 (15.3) | 37 (18.4) | 0.626 | 35 (19.2) | 17 (14.5) | 0.349 |
| 1-year mortality | 89 (29.8) | 24 (24.5) | 65 (32.3) | 0.180 | 60 (33.0) | 29 (24.8) | 0.154 |
| Refracture rate | 8 (2.7) | 2 (2.0) | 6 (3.0) | >0.999 | 6 (3.3) | 2 (1.7) | 0.488 |
| 30-day readmissions | 21 (7.0) | 6 (6.1) | 15 (7.5) | 0.811 | 13 (7.1) | 8 (6.8) | >0.999 |
| 1-year readmissions | 72 (24.1) | 20 (20.4) | 52 (25.9) | 0.317 | 44 (24.2) | 28 (23.9) | >0.999 |

| Risk factor, n (%) | Alive (<i>n</i> =210) | Died (n=89) | <i>p</i> -value |
|--------------------|------------------------|-------------|-----------------|
| Age group, years | | | |
| <65 | 58 (89.2) | 7 (10.8) | < 0.001 |
| ≥65 | 152 (65.0) | 82 (35.0) | |
| Sex | | | |
| Male | 62 (65.3) | 33 (34.7) | 0.222 |
| Female | 148 (72.55) | 56 (27.45) | |
| Fracture type | | | |
| NOF | 104 (68.0) | 49 (32.0) | 0.448 |
| Pertrochanteric | 106 (72.6) | 40 (27.4) | |
| ASA category | | | |
| 1 or 2 | 118 (83.1) | 24 (16.9) | < 0.001 |
| ≥3 | 92 (58.6) | 65 (41.4) | |

| | | Group | | | Group | | |
|---|-----------|-----------------|--------------------|-----------------|-------------|-----------|---------|
| Variable, n (%) | All | Bisphosphonates | No bisphosphonates | <i>p</i> -value | Pre-FLS era | FLS era | p-value |
| Neck of femur fracture, <i>n</i> =153 | 153 | 46 | 107 | | 97 | 56 | |
| 30-day mortality | 16 (10.5) | 5 (10.9) | 11 (10.3) | >0.999 | 10 (10.3) | 6 (10.7) | >0.999 |
| 90-day mortality | 27 (17.6) | 10 (21.7) | 17 (15.9) | 0.488 | 15 (15.5) | 12 (21.4) | 0.383 |
| 1-year mortality | 49 (32.0) | 14 (30.4) | 35 (32.7) | 0.852 | 31 (32.0) | 18 (32.1) | >0.999 |
| Refracture incidence | 6 (3.9) | 1 (2.2) | 5 (4.7) | 0.669 | 5 (5.2) | 1 (1.8) | 0.416 |
| 30-day readmissions | 9 (5.9) | 3 (6.5) | 6 (5.6) | >0.999 | 5 (5.2) | 4 (7.1) | 0.725 |
| 1-year readmissions | 40 (26.1) | 10 (21.7) | 30 (28.0) | 0.548 | 25 (25.8) | 15 (26.8) | >0.999 |
| Pertrochanteric femur fracture, <i>n</i> =146 | 146 | 52 | 94 | | 85 | 61 | |
| 30-day mortality | 12 (8.2) | 4 (7.7) | 8 (8.5) | >0.999 | 8 (9.4) | 4 (6.6) | 0.761 |
| 90-day mortality | 25 (17.1) | 5 (9.6) | 20 (21.3) | 0.107 | 20 (23.5) | 5 (8.2) | 0.024 |
| 1-year mortality | 40 (27.4) | 10 (19.2) | 30 (31.9) | 0.122 | 29 (34.1) | 11 (18.0) | 0.039 |
| Refracture incidence | 2 (1.4) | 1 (1.9) | 1 (1.1) | >0.999 | 1 (1.2) | 1 (1.6) | >0.999 |
| 30-day readmissions | 12 (8.2) | 3 (5.8) | 9 (9.6) | 0.539 | 8 (9.4) | 4 (6.6) | 0.761 |
| 1-year readmissions | 32 (21.9) | 10 (19.2) | 22 (23.4) | 0.677 | 19 (22.4) | 13 (21.3) | >0.999 |

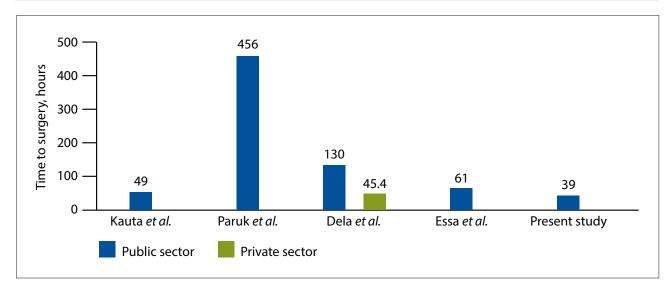


Fig. 1. Time to surgery (hours) in South African studies. $^{[1,30,35,36]}$

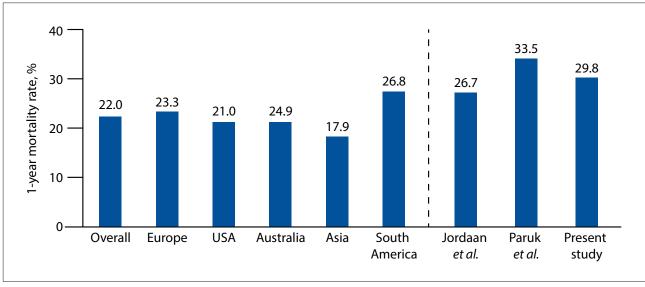


Fig. 2. One-year mortality rate comparison from the studies by Downey et al., [39] Paruk et al. [35] and Jordaan et al. [40]

in SSA poses a significant challenge to meeting these goals. [6] Moreover, SA healthcare is currently based on an unbalanced two-tiered medical system, where only ~16% of the population have private medical insurance. The orthopaedic surgeon density per 100 000 uninsured population is 0.36, compared with the private sector at 8.3, and World Health Organization target of >5.[28,29] Access to appropriate care is therefore often limited in public hospitals.^[29,30] Even when certain standards are met, the evaluation and management of osteoporosis and orthopaedic-geriatric comanagement are notoriously neglected, both locally and internationally.[1,29,31-34]

In the current study, the median time to surgery following an FFH was 39.0 hours, which compares favourably with other local research,[1,35,36] and was, surprisingly, even quicker than what was reported from private sector hospitals in the 2022 study by Dela et al.[30] (Fig. 1). It is worth noting that after implementation of the FLS, the time from diagnosis to surgery further improved from 42.5 hours to 30.8 hours. While the FLS did not include implementation of an accelerated time to surgery, it is tempting to speculate that the introduction of an FLS might have led to behavioural changes in the orthopaedic department due to an improved understanding of 'priority' of these patients. The time to surgery was calculated from the time that a diagnosis was made to the 'cutting time' in theatre, meaning that in the 73.2% of patients who were referred from outside facilities, this time included ambulance transport.

During the FLS era, 83.8% of FFH patients were initiated on bisphosphonate therapy, calcium and vitamin D supplementation, and referred to internal medicine and endocrine specialist units for continuation of multidisciplinary care, representing a marked improvement from the era before the FLS.

The 30-day, 90-day and 1-year all-cause mortality rates were 9.4%, 17.4% and 29.8%, respectively. Differences observed in 1-year mortality between the FLS period and the pre-FLS period, or in the bisphosphonate cohort v. non-bisphosphonate cohort, did not reach statistical significance in our study. The minor differences observed during the subanalyses, although interesting to note, should be interpreted with caution, as the authors are unaware of any prior literature suggesting superior efficacy of bisphosphonates or FLS for specific fracture types or age groups. These findings, however, raise interesting questions that future studies may investigate.

While international mortality rates are reported to have largely stayed constant over a period of three to four decades,[37,38] a large systematic review in 2019 by Downey et al.[39] suggests that 1-year mortality may now be slowly declining globally, but remains >20% in most countries outside of Asia. Other SA studies have reported 1-year mortality rates of 33.5%^[35] and 26.7%,^[40] although the latter was limited to fractures treated with arthroplasty (Fig. 2).

Unplanned hospital readmission is an undesirable complication associated with worse outcomes, including higher 1-year mortality. [41] In the present study, readmission occurred at a frequency of 7.0% at 30 days, and 24.1% within the first year. These figures compare favourably with previous literature. Data from the USA suggest a rate of 11.5% at 30 days, [42] and a recent Swedish study reported a 1-year readmission rate of 39%.[43] Both these studies were large-scale, based on national hip fracture registries.

Refractures within the first year occurred in 2.7% (n=8) of patients in the present study. This was lower than in a Spanish study that reported a 4.1% first-year refracture rate, [12] as well as in the landmark Horizon trial by Lyles et al., [15] which reported 8.6% in the bisphosphonate group and 13.9% in the placebo group. Of note, the latter study had a longer follow-up and did not report the 1-year refracture specifically. It is possible that some patients in the present study may have sustained secondary fractures, and either

never presented, or presented to health facilities outside the province. However, these numbers are likely very low.

The low refracture and unplanned readmission rates may reflect good perioperative care, a population with inherent resilience and/or social support structures, or difficulties in accessing health facilities. The last, however, is a less likely explanation, as the low refracture and readmission rates are not offset by higher-than-average mortality rates. As expected, older age and a higher ASA score are reported as risk factors for mortality at 1 year following surgery.

Study strengths and weaknesses

This study contributes to a growing body of evidence surrounding FFH in SSA. To the authors' best knowledge, it is the first local study to specifically evaluate outcomes within an FLS, and as the incidence will continue to increase, it may assist in informing health policymakers and clinicians. It also serves as an example to other local hospitals setting up an FLS, showing that it can be successfully done at a regional level.

This study has several limitations. No data were collected on coronavirus infections, even though the study coincided almost perfectly with the COVID-19 pandemic and its sequential waves in SA. The effects thereof, although not underestimated, are difficult to quantify. It is known that FFH patients infected with COVID had higher 30-day mortality rates, but there was no increased mortality in uninfected FFH patients during the pandemic. [44,45] There is an element that is system and time specific: the lower trauma burden during the same period could have facilitated a lower ratio of patients to health providers in an orthopaedic unit, leading to more efficient perioperative care.

The retrospective nature of the study has inherent limitations, especially considering data collection. Of note, good record-keeping, availability of multiple electronic databases and collaboration with the Department of Home Affairs greatly facilitated both collection and verification of accurate data. The sample size was relatively small (n=299), and the follow-up was only 1 year. In addition, imbalances in study groups are common limitations in retrospective cohort studies. In this study, however, all baseline characteristics were balanced across study groups.

Patients were sampled from a single centre in a single province, which means that the ethnic composition of patients may not reflect that of the whole country. Caution should be taken when generalising these results to the rest of SA, or other countries, as it is known that there is a geographical and ethnic influence on incidence as well as outcomes after FFH.[6]

Given the small proportion of patients who died within 1 year (29.8%), taken together with the relatively small total cohort size, the study was powered to detect only large differences between groups. Therefore, future local studies should attempt to follow patients for longer periods or potentially include multiple sites, to increase the overall sample size and allow for the detection of risk factors for mortality in a SA context.

Conclusion

Successful implementation of an FLS in a regional SA hospital did not translate to significant reduction in early mortality, refractures or readmissions. However, it resulted in a marked increase in bisphosphonate initiation, and a reduction in time to surgery.

Data availability. The data used for this study are available from the authors on request.

Declaration. This study was done in partial fulfillment of SPO's MMed (Orthopaedic Surgery) degree at Stellenbosch University.

Acknowledgements. Dr Gerhard Thiart - Worcester Provincial Hospital orthopaedic department.

Author contributions. SPO: design of study protocol, data collection, writing of manuscript. MCB: study design, data analysis, supervision, editing. JDJ: study conceptualisation, supervision, editing.

Funding. None.

Conflicts of interest. None.

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Received 15 January 2025; accepted 4 June 2025.