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Endoprostheses Last Longer Than Intramedullary Devices in Proximal Femur Metastases

Norah Harvey MD, Elke R. Ahlmann MD, Daniel C. Allison MD, MBA, Lingjun Wang PA, Lawrence R. Menendez MD, FACS

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Abstract

Background The proximal femur is the most common site of surgery for bone metastases, and stabilization may be achieved through intramedullary fixation (IMN) or endoprosthetic reconstruction (EPR). Intramedullary devices are less expensive, less invasive, and may yield improved function over endoprostheses. However, it is unclear which, if either, has any advantages.

Questions/purposes We determined whether function, complications, and survivorship differed between the two approaches.

Methods We retrospectively reviewed 158 patients with 159 proximal femur metastatic lesions treated with surgical stabilization. Forty-six were stabilized with IMN and 113

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Each author certifies that his or her institution has approved the human protocol for this investigation, that all investigations were conducted in conformity with ethical principles of research, and informed consent for participation in the study was obtained.

N. Harvey, E. R. Ahlmann, D. C. Allison, L. Wang, L. R. Menendez
Department of Orthopedics, University of Southern
California + Los Angeles County Medical Center,
Los Angeles, CA, USA

D. C. Allison (⋈)
Department of Orthopedics, Division of Musculoskeletal
Oncology, Keck School of Medicine, USC University
Hospital, Los Angeles, CA, USA
e-mail: dallison@usc.edu

L. R. Menendez Department of Orthopedics, Keck School of Medicine, USC University Hospital, Los Angeles, CA, USA Introduction

One-third of metastatic lesions to the skeleton occurs in the proximal femur [11]. Approximately 10% of patients with metastatic disease will sustain pathologic fractures, and 65%

were treated with EPR. The minimum followup was 0.25 months (mean, 16 months; median, 17 months; range, 0.25–86 months).

Results The mean Musculoskeletal Tumor Society score was 24 of 30 (80%) after IMN and 21 of 30 (70%) after EPR. There were 12 complications (26%) in the IMN group, including 10 nonunions, six of which went on to mechanical failure. There were complications in 20 of 113 (18%) of the EPR group, which consisted of 10 dislocations (9%) and 10 infections (9%). There were no mechanical failures with EPR. Both implants remained functional for the limited lifespan of these patients in each group at all time intervals. EPRs were associated with increased implant longevity compared with IMNs (100% versus 85% 5-year survival, respectively) and a decreased rate of mechanical failure (0% versus 11%, respectively) when compared with the intramedullary devices.

Conclusions Patients with metastatic disease to the proximal femur may live for long periods of time, and these patients may undergo stabilization with either IMN or EPR with comparable functional scores and the implant survivorship exceeding patient survivorship at all time intervals. Endoprostheses demonstrate a lower mechanical failure rate and a higher rate of implant survivorship without mechanical failure than IMN devices.

Level of Evidence Level III, therapeutic study. See Guidelines for Authors for a complete description of levels of evidence.



of all fractures requiring surgery occurs in the femur [11, 12]. Impending and pathologic fractures of the proximal femur can have a disastrous effect on the patient's quality of life, resulting in severe pain and limitation in function and mobility. Current estimates state that 40% of patients with pathologic fractures will survive for at least 6 months after their fracture and 30% will survive for more than 1 year [16]. Patients with metastatic disease are living longer, increasing the risk of failure of conventional techniques.

The primary goals of surgical stabilization of proximal femur metastatic lesions are to relieve pain, allow immediate return to function, improve the patient's quality of life, facilitate medical testing and treatments, and decrease the chance of subsequent surgical revision. Surgical options that have emerged to achieve these goals include plate fixation [13, 27], intramedullary fixation [6, 13, 24, 26, 27], and endoprosthetic reconstruction [4, 5, 19, 28-30]. Plate fixation has an unacceptably high mechanical failure rate of 44% to 70% [6, 13, 30]. Intramedullary devices have a lower rate of mechanical failure of 2% to 22% but death from cardiopulmonary complications ranges from 1% to 10% [6, 24, 26, 28]. Endoprostheses have the lowest reported mechanical failure rate (under 3.7%) with a complication rate of 6% to 35% [5, 21, 25]. One study directly comparing the use of intramedullary devices and endoprostheses for proximal femur metastatic lesions found an increased 2-year risk of reoperation when comparing internal fixation with endoprosthetic reconstruction [28]. However, it is unclear whether function, complications, or implant survivorship differs between the two treatments.

We therefore compared (1) function; (2) complications; and (3) survivorship in patients treated with intramedullary devices and endoprosthetic reconstruction.

Patients and Methods

From our institution's Orthopaedic Oncology Database we identified all 167 patients with metastatic disease of the proximal femur who underwent surgery with either intramedullary nailing or endoprosthetic reconstruction from 1998 to 2009. We included all patients with lesions of the proximal one-third of the femur who underwent either

reconstruction nailing or resection of the proximal femur with placement of an endoprosthesis. Lesions located only in the femoral head without peritrochanteric extension were excluded as were patients with a lesion not amenable to either treatment method or who underwent revision of a preexisting implant. Indications for surgery were (1) an impending (Mirels score of 9 or greater [18]) or existing pathologic fracture; and (2) intractable pain and loss of ambulatory ability. Contraindications to surgery included (1) projected survival of less than 6 weeks; and (2) lack of medical fitness to undergo the procedure. After nine patients were excluded, 158 patients with 159 cases were available for review. There were 72 males and 86 females with a mean age of 60 years (range, 17-91 years) (Table 1). Impending fractures accounted for 43% (68 of 159) and actual pathologic fractures made up 53% (91 of 159) of the total lesions treated (Table 1). Diagnoses were comparatively distributed with breast adenocarcinoma (26% [41 of 158]) as the most common overall (Table 2). Thirty-four of 158 (22%) patients were alive at the time of this study. Of the patients treated with intramedullary

Table 2. Distribution of diagnoses

Diagnosis	Total (%)	Reconstruction nail (%)	Endoprosthesis (%)
Breast	41 (26)	10 (22)	31 (27)
Sarcomas	20 (13)	12 (26)	8 (7)
Lung	19 (12)	2 (4.3)	17 (15)
Renal	19 (12)	5 (11)	14 (12)
Multiple myeloma	14 (9)	9 (20)	5 (4.5)
Prostate	10 (6)	1 (2.2)	9 (8)
Colorectal	7 (4.4)	4 (6.5)	3 (2.7)
Melanoma	6 (3.8)	2 (4.3)	4 (3.6)
Thyroid	4 (2.5)	0	4 (3.6)
Liver	4 (2.5)	1 (2.2)	3 (2.7)
Cervical/uterine	4 (2.5)	2 (4.3)	2 (1.8)
Bladder	4 (2.5)	1 (2.2)	3 (2.7)
Unknown	3 (1.9)	0	3 (2.7)
Lymphoma	2 (1.3)	2 (4.3)	0
Neuroendocrine	1 (0.6)	1 (2.2)	0
Total	158 (100)	45 (28)	113 (72)

Table 1. Patient demographics in regard to type of surgical treatment

Surgical treatment	Total number (M/F)	Average age (years)	Average followup (months)	Fracture	
				Impending (% of total)	Pathologic (% of total)
Endoprosthesis	113 (53 M/60 F)	61 (range, 17–86)	14 (range, 0.75–86)	43/113 (38%)	70/113 (63%)
Reconstruction nail	46 (19 M/26 F)	56 (range, 16–91)	20 (range, 0.25–78)	25/46 (54%)	21/46 (46%)

M = male; F = female.

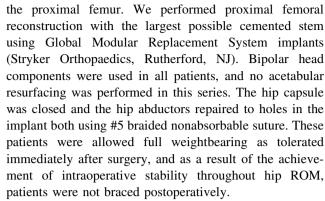


fixation, 54% (25 of 46) had impending fractures and 46% (21 of 46) had existing fractures of the proximal femur; of the patients treated with endoprosthetic reconstruction, 38% (43 of 113) had impending fractures and 62% (70 of 113) had pathologic fractures (Table 1). The minimum patient followup from the time of surgery was 0.25 months (average, 16 months; range, 0.25–86 months) (Table 1) with no patients lost to followup. Thirty-five patients were alive at the time of this review (15 in the intramedullary fixation [IMN] group and 20 in the endoprosthetic reconstruction [EPR] group) with the minimum followup of these patients being 24.4 months (median, 17 months; range, 9–78 months). No patients were recalled specifically for this study; all data were obtained from medical records and radiographs, which were complete in all cases. This study was approved by our institution's investigational board review.

The published mechanical failure rate of intramedullary devices has been roughly 20%, whereas that of EPR approximates 0% [5, 6, 21, 24–26, 28, 29]. Therefore, we wanted to detect at least a 20% to 25% higher failure rate of the intramedullary devices. A higher failure rate would be clinically unacceptable. We performed a power analysis sample size calculation for 25% IMN failure and 1% endoprosthetic failure and concluded that a sample size of 52 in each group will have 80% power to detect a 24% difference using a two-group t-test with a 0.050 two-sided significance level.

We presented patients with both treatment options, counseled about the risks and benefits of each, and allowed them to choose their preferred method of treatment. Patients were encouraged to undergo resection and EPR in the presence of an existing fracture, more than 50% canal involvement on plain films, or isolated renal cell or thyroid carcinoma. We performed IMN with either the TriGen Femoral Antegrade Nail in reconstruction locking mode (Smith and Nephew Inc, Memphis, TN) or the long Titanium Trochanteric Fixation Nail System (Synthes Inc, West Chester, PA) using the manufacturers' recommended technique. The goal was to insert the largest possible diameter nail without reaming the femur to limit further spread of disease both locally and systemically. For patients with lesions involving greater than one-third of the diameter of the bone or greater than 2.5 cm in cortical destruction, we performed curettage and placement of supplemental cement for additional structural support. Postoperatively, patients with pathologic fractures treated with IMN were kept partially weightbearing until there was radiographic evidence of healing of at least two cortices. All other patients were allowed immediate weightbearing as tolerated after the procedure.

All proximal femur EPRs were performed using a standardized protocol through a posterolateral approach to



Patients from the intramedullary fixation group received a standardized protocol of 30 Gray of external beam radiation starting 3 weeks after surgery. Both groups resumed their chemotherapy as indicated by their disease.

Routine followup examinations were performed at 1 week, 4 weeks, 2 months, 6 months, 1 year, and yearly thereafter. Charts and radiographs were available for review for all patients in this study, and recorded outcomes were based on the most recent followup evaluation. The Musculoskeletal Tumor Society (MSTS) functional scale for the lower extremity (which specifically evaluates supports, walking ability, and gait) [7] was administered by an investigator (LW) not involved in patient care to avoid potential bias. Complications were defined as adverse events that required additional surgical intervention; we did not consider other complications in this analysis. Charts were reviewed for the onset of complications, including infection, recurrence, aseptic loosening, fatigue failure, periprosthetic fracture, and dislocation.

Radiographs taken at the latest followup were evaluated for evidence of hardware failure or loosening, periprosthetic fracture, and nonunion. One investigator (ERA) evaluated preoperative radiographs for location of the lesion and percent of canal involvement as well as most recent followup radiographs for complications. Loosening was determined radiographically by the presence of circumferential radiolucency of greater than 2 mm around the implant in any zone and implant subsidence/migration of any distance in regard to the position of the implant stem to the surrounding cortical bone [9, 20]. Radiographic nonunion was defined as failure of a fracture to unite radiographically (bridging of four cortices on two orthogonal radiographs) with associated pain by 6 months from the date of fixation. Complication-free survival was defined as survivorship of the patient without any complications. Implant survivorship was defined as percentage of functional implants remaining that did not undergo revision of any or all components or removal of implants without revision. Survivorship of the implant without mechanical failure describes implant survival without experiencing a structural mechanical failure of the actual implant itself, ie, breakage of the nail, breakage of



the stem, disassembly of the bipolar components, breakage of the nail's interlocking bolts, etc.

We determined differences in MSTS scores between the two groups using a t-test. Differences in complications were determined using Fisher's exact test and Kaplan–Meier survivorship curves using patient death and failure of the reconstruction as end points and corresponding 95% confidence intervals were generated using GraphPad Prism® software (GraphPad Software, Inc, San Diego, CA). We used log-rank analysis to determine differences in survival curves. Survivorship data met assumptions associated with the statistical analysis. The t-test comparison of MSTS score means agreed with our assumptions of continuous data, which followed a normal distribution, reflecting a random sample from the population. We used SAS/STAT® software (SAS® Institute, Inc, Cary, NC) for all analyses for all comparisons.

Results

The MSTS functional scores were similar (p = 0.28) for living patients treated with an intramedullary device and

those treated with EPR: 80% (24 of 30; range, 8–30) versus 70% (21 of 30; range, 12–27), respectively.

We found no difference in the rate of adverse events requiring surgery (p = 0.097) or the infection rate (p =0.283) between patients treated with an endoprosthesis and those treated with an intramedullary nail. We observed more complications (unrelated to recurrence or local disease) 1 year after surgery (p < 0.001) in patients treated with intramedullary fixation compared with endoprostheses: 67% versus 6%, respectively. There were 12 (26%) complications requiring further surgery in patients having intramedullary fixation. These included 10 (22%) symptomatic nonunions, of which six (13%) went on to have mechanical failure of the implant, one experienced deep infection, and one painful hardware (Table 3). We found no difference in the failure rate of the intramedullary devices based on location of the lesion (p = 0.723) or percent of canal involvement (p = 0.737). There were no failures in patients with greater than twothirds canal involvement. All mechanical failures occurred in patients with one-third to two-thirds canal involvement. IMN cases with actual fractures had a higher rate of both mechanical failure (p = 0.028) and

Table 3. Intramedullary nail complications

Age (years), gender	Primary tumor	Complication	Time to complication	Treatment	Outcome	Followup
46, M	Liposarcoma	Nonunion and nail breakage	78 months	Removal of nail; placement of bipolar endoprosthesis	No further complications	4 months
16, F	Glomangio- sarcoma	Nonunion and nail breakage	14 months	Removal of nail; placement of bipolar endoprosthesis	No further complications	45 months
60, F	Breast	Nonunion and nail breakage	18 months	Removal of nail; placement of bipolar endoprosthesis	No further complications	52 months
60, M	Renal	Nonunion and nail breakage	11 months	Removal of nail; placement of bipolar endoprosthesis	No further complications	19 months
80, M	Renal	Nonunion and nail breakage	19 months	Removal of nail; placement of bipolar endoprosthesis	No further complications	27 months
50, F	Breast	Nonunion and breakage of both distal interlocking screws	6 months	Removal of nail; placement of bipolar endoprosthesis	No further complications	8 months
50, F	Sarcoma	Nonunion	61 months	Revision to larger diameter nail; bone graft	Persistent nonunion	3 months
68, F	Sarcoma	Nonunion	14 months	Revision to larger diameter nail; bone graft	Developed deep infection 4 months postoperatively; hip disarticulation	36 months
51, F	Sarcoma	Nonunion	5 months	Revision to larger diameter nail	Union of fracture at 6 months	19 months
40, F	Sarcoma	Nonunion	16 months	Revision to larger diameter nail	Union of fracture at 6 months	22 months
54, M	Sarcoma	Infection	4 months	Hip disarticulation	No infection	23 months
31, F	Lymphoma	Painful hardware (distal interlocks)	22 months	Removal of distal interlocks	No further complications	46 months

M = male; F = female.



reoperation (p = 0.011) than those with impending fractures. Complications in patients treated with endoprostheses included 10 dislocations (9%) and 10 deep infections (9% [10 of 113]) (Table 4). We were unable to find a difference in the overall complication rate based on pathologic diagnosis. There was a lesser rate of mechanical failure in the endoprosthesis group compared with the intramedullary nail group (p = 0.003). There were no intraoperative deaths in this series and no patients treated with EPR died within 14 days of surgery, whereas only one patient who underwent intramedullary stabilization died from pneumonia and sepsis during this time.

The overall patient survival in this series was 51%, 29%, and 11% at 1 year, 2 years, and 5 years, respectively. According to Kaplan-Meier analysis, the survival of both the intramedullary implants (p = 0.008, log-rank test) and the endoprostheses (p = 0.001) at 5 years exceeded that of the patients, a finding that was also unrelated to recurrence or local disease (Fig. 1). We found no difference (p = 0.306) in complication-free survival of the patients in either group (Fig. 2). However,

the endoprosthesis group demonstrated increased survival of the implant without mechanical failure with 100% 2- and 5-year survival compared with 85% 2- and 5-year survival of the intramedullary fixation group (p = 0.003) (Fig. 3).

Overall Patient Survival Versus Implant Survival

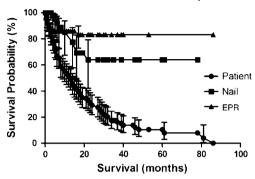


Fig. 1 This Kaplan–Meier survivorship curve with 95% confidence intervals shows that both nails (p=0.008, log-rank test) and endoprostheses (EPR) (p=0.001, log-rank test) outlived the patients in whom they had been implanted.

Table 4. Endoprosthetic reconstruction complications

Age (years), gender	Primary tumor	1		Treatment	Outcome	Followup
53, F	Breast	Dislocation	1.5 months	Open reduction	Stable	2 months
		Infection	2 months	I&D, IV Abx	No infection	
68, M	Prostate	Dislocation and infection	1 month	I&D, resection arthroplasty, IV Abx	No infection, no reimplantation	21 months
52, F	Breast	Dislocation	3 months	Open reduction	Stable	9 months
		Infection	4 months	I&D, IV Abx	No infection	
67, F	Liposarcoma	Dislocation	3 months	Open reduction	Stable	6 months
		Infection	5 months	Hip disarticulation	No infection	
55, F	Sarcoma	Infection	3 months	I&D, IV Abx	No infection	29 months
62, M	Lung	Infection	3 months	I&D, IV Abx	No infection	6 months
61, F	Breast	Infection	1 month	I&D, IV Abx	No infection	3 months
73, M	Lymphoma	Infection	17 months	I&D, IV Abx	No infection	19 months
74, M	Thyroid	Infection	2 months	I&D, IV Abx	No infection	9 months
28, M	Hepatic	Infection	1 month	I&D, IV Abx	No infection	6 months
44, F	Melanoma	Dislocation	1 month	Open reduction	Stable	4 months
68, M	Lung	Dislocation	1 month	Closed reduction	Stable	2 months
82, M	Renal	Dislocation	2 months	Closed reduced	Stable	81 months
68, F	Breast	Dislocation	1 month	Open reduction, lengthening of intercalary parts for more stability	Stable	29 months
67, M	Melanoma	Dislocation	1 month	Open reduction	Stable	2 months
59, F	Breast	Dislocation	2 months	Open reduction	Stable	10 months

F = female; M = male; I&D = irrigation and débridement; IV Abx = intravenous antibiotics.



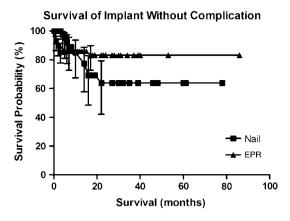


Fig. 2 This Kaplan–Meier survivorship curve with 95% confidence intervals shows that there was no difference (p = 0.31) in complication-free survival between nails and endoprostheses (EPR).

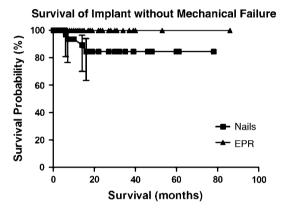


Fig. 3 This Kaplan–Meier survivorship curve with 95% confidence intervals shows improved survival of the implant without mechanical failure in the endoprosthesis (EPR) group (p = 0.003).

Discussion

Surgical management of patients with metastatic disease of the proximal femur has evolved over the last 30 years with a move away from plate fixation to the use of intramedullary and endoprosthetic devices. However, the ideal method of treatment still remains unclear. We therefore compared (1) function; (2) complications; and (3) survivorship in patients treated with intramedullary devices and EPR.

We acknowledge several limitations of this study. First, owing to the nonrandomized retrospective design, we did not control for potentially confounding variables such as type or extent of disease, the degree of intraoperative resection, preoperative level of function, age of the patient, medical problems (osteoporosis), and associated adjuvant treatments (chemotherapy, bisphosphonates, etc). Metastatic lesions to the proximal femur occur in a very diverse group with a variety of diagnoses at different stages of disease and with differing lesion characteristics. Each

patient's treatment must be individualized to their current situation, making it difficult to conduct a truly randomized study comparing one surgical technique with another with sufficient numbers. However, all patients were treated at one institution using the same rationale, selection criteria, and surgical technique. Second, there is selection bias in the choice of surgical implant. When discussing surgical options with our patients, we presented risks and benefits of both IMN and EPR, allowing the patient to make an informed decision regarding implant choice based on their personal needs and expectations. For patients with extensive lesions (involving greater than 50% of the bone diameter) and fractures, we generally encouraged resection and EPR; however, not all patients chose this option. Even so, this bias would skew the endoprosthetic group toward more aggressive tumors and extensive lesions. Nonetheless, we observed no difference in overall complication rates but rather a decreased rate of mechanical failure (0% versus 11%) in the endoprosthetic group. Selection bias will be difficult to eliminate from future studies, because each patient will continue to be treated on an individual basis. Third, our complication rate only measured those complications requiring surgical intervention, which may exclude nonsurgical events (excessive bleeding, nerve palsy, chronic pain, etc) of clinical significance and should be noted. However, the objective nature of this measure lends itself to more accurate reporting of complications as opposed to the subjective nature of the measures "minor" or "major." Next, we treated all patients with failed implants with surgical revision, although not all patients with metastatic disease will have revision surgery. This difference may skew functional results when compared with patients outside of this study. Patients with metastatic disease to bone have a decreased life expectancy; therefore, survivorship analysis may not accurately reflect the true longevity of the implant in all patients. However, the study specifically evaluates patients with metastatic disease and therefore should accurately reflect survivorship in the cohort being treated.

Our MSTS scores of 80% of normal in the intramedullary nail group and 70% of normal in the endoprosthesis group compare with scores reported by Potter et al. in patients with metastatic disease to the proximal femur treated with EPR [22]. Our MSTS score percentages of normal also compare with a Toronto Extremity Salvage Score of 61% of normal reported by Chandrasekar et al. in their series of proximal femoral replacement [5].

We found no difference in complications between our two groups. In our intramedullary fixation group, 22% (10 of 46) experienced painful nonunion, of which 60% (six of 10) of cases eventually went on to hardware failure. The presence of fracture resulted in an increased rate of reoperation and subsequent mechanical failure in the



intramedullary nail group. Implant failure has been reported in other series evaluating IMN for metastatic lesions with a failure rate ranging between 2% and 22% [13, 26–28]. This probability of implant failure increases with longer patient survival and the presence of nonunion [6]. Intramedullary nails are designed to act as internal splints with load-sharing properties and are assumed to bear most of the load initially and then gradually transfer it to the bone as the fracture heals [3]. These devices were not designed to bear the entire load of the patient for the remainder of the patient's lifetime and are thus at a higher risk for failure in patients with large bone defects or fractures may never heal. Furthermore, intralesional treatment of metastatic disease is treated with adjuvant external beam radiation therapy, which also may contribute to delayed union and resultant implant failure. In contrast, the loadbearing characteristics of endoprostheses allow immediate postoperative stability and weightbearing with a low reported mechanical failure rate [4, 5, 25, 28]. We found endoprostheses had a lower rate of mechanical failure than intramedullary devices (0% versus 11%). Another study that directly compared these two groups found a statistically insignificant trend toward decreased mechanical failure in the endoprosthesis group [28]. A majority of the intramedullary nail complications occurred late (after 1 year) as compared with a small minority of the endoprosthesis complications, a comparison not yet noted in previous literature but which should play a role in decisionmaking. Our relatively low dislocation rate of 9% (10 of 113) is comparable to previous series of endoprostheses using bipolar heads, which demonstrate a 1% to 12% dislocation rate [4, 5, 10, 21, 23, 25]. Other series in which acetabular resurfacing was also performed report a higher rate of instability of up to 30% when compared with bipolar implants [10, 17]. Dislocation may be further reduced by performing capsular repair [17]. It is our recommendation to use bipolar head components and repair the hip capsule whenever possible. Previous studies cite perioperative cardiopulmonary complications attributed to the use of long intramedullary implants with a rate of perioperative death as high as 10% [2, 14, 19, 25]. In our study, one patient died within 2 weeks of IMN and none within 14 days of EPR. We attribute our lack of mortality to our preoperative medical optimization, use of unreamed femoral nails, use of sharp reamers for long-stem endoprostheses, and careful cementing technique, allowing the anesthesiologist to optimize the patient's vital parameters and increase intravenous fluids before cementing.

In a previous series of 228 pathologic long bone fractures, the most important risk factor for treatment failure was the length of survival after surgery [6]. Because patients are living longer with improved adjuvant therapies, this treatment failure is expected to further increase. Some

authors argue that, for patients with a long projected survival, an endoprosthesis is the more durable surgical option [5, 21]. Some have even suggested that EPR should be the treatment of choice in isolated lesions from renal cell metastasis because these patients may have prolonged survival [1, 15]. In our series, 51% of patients survived 1 year, 29% survived 2 years, and 11% survived 5 years. Our endoprosthesis survivorship compares well with that reported in of the literature, which has demonstrated 86% to 93% revision-free implant survival [5, 8, 22]. Although we found no difference in complication-free survival between our two patient groups, we found higher implant survival without mechanical failure in the endoprosthesis group, which supports data showing increased mechanical failure among intramedullary devices.

When treating metastatic lesions to the proximal femur, both IMN and EPR may be used for stabilization with similar functional scores and overall complication rates, and both devices generally outlasted the patient. Patients with metastatic disease to the bone are living longer than previously considered, and surgery should be performed with this extended survival in mind. Intramedullary fixation in the presence of fracture leads to increased mechanical failure and reoperation rates, and most of the intramedullary complications occur after 1 year as opposed to EPR complications, which occur early. Advantages of EPR over IMN include the immediate ability to fully weightbear, the avoidance of nonunion, and the decreased incidence of mechanical failure and subsequent revision surgery. Therefore, we believe EPR should be considered in patients with proximal femur metastatic lesions with associated pathologic fracture and prolonged life expectancy.

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