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*2015 Annual Meeting*

*Sports Medicine/Pediatrics/Tumor  
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Kiawah Island, South Carolina

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and the Southern Regional Area Health Education Center

## Evaluating the Effect of an Off-the-Shelf Hip Orthosis on Balance in Post-operative Hip Arthroscopy: A Pilot Study

Benjamin M. Wooster, MD<sup>1</sup>, Elizabeth A. Howse, MD<sup>2</sup>  
Benjamin L. Long, MS<sup>3</sup>, Judy Foxworth, PT, PhD<sup>3</sup>,  
Allston J. Stubbs MD, MBA<sup>4</sup>

1. Duke University Medical Center, Dept. of Orthopaedic Surgery, Durham, NC 2.  
Long Island Jewish Medical Center, Dept. of Emergency Medicine, New Hyde Park, NY 3. Winston-Salem State University, Dept. of Physical Therapy, Winston Salem, NC  
4. Wake Forest Baptist Health, Dept. of Orthopaedic Surgery, Winston-Salem, NC



## Disclosure

- Dr. Stubbs has financial relationships with the following companies:
  - Consultant: Smith & Nephew
  - Stock: Johnson & Johnson
  - Research Support: Bauerfeind
  - Department Support: Smith & Nephew Endoscopy, Depuy, Mitek
  - Boards/Committees: AOSSM, ISHA, AANA
- All other authors report no declarations of interest
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## Background

- Hip orthoses commonly utilized in post-op rehab
- Primary functions
  - Restrict range of motion
  - Protect compromised tissue
  - Role in balance?
- Limitations in literature
  - Variations in post-op rehab protocols
  - Lack of randomized control trials
  - Expert opinion

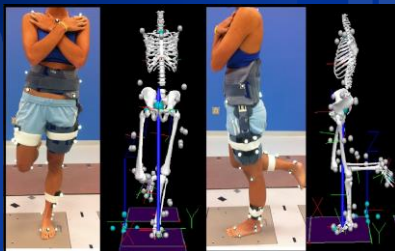
## Methods

- Patient Selection
  - Wake Forest Baptist Medical Center
  - Hip arthroscopy for pathology associated with FAI
- Data Collection
  - WFU Human Performance and Biodynamics Laboratory
  - Four weeks post-operative appointment
- Off-The-Shelf Hip Orthosis
  - Sof-Tec Coxa®, Bauerfeind AG, Zeulenroda, Germany



## Methods

- Barefoot static single leg postural sway test on force plate for 60 second trials

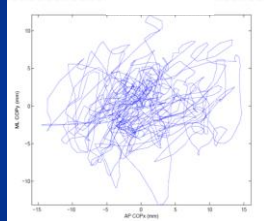


## Methods

- Initial testing side and bracing status randomized
- Unbraced trials
  - Alternated between legs
  - Separated by at least one minute of rest
- Braced trials
  - Limited to one leg
  - Separated by at least two minutes of rest
- Three valid trials recorded for each condition *or* until six unsuccessful attempts per condition
- Two best trials included in final data analysis

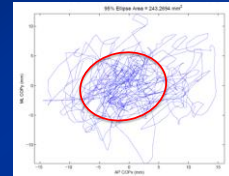
## Methods

- MatLAB® Software
  - Mathworks, Natick, MA
- Center of pressure trajectory
  - Ground reaction forces
    - $F_x$ ,  $F_y$ ,  $F_z$
  - Ground reaction moments
    - $M_x$ ,  $M_y$ ,  $M_z$
- Stabilograms



## Methods

- Center of Pressure Ellipse Area (COPEA)
  - Ellipse encircling 95% of the data points
  - Anterior-posterior and medial-lateral motion



*Increases in COPEA = Decreased balance*

## Results: Demographics

	IMPROVED (N=10)	WORSENE (N=7)	P
Sex	9F / 1M	7F / 0M	
Age (yrs)	25.27 ± 5.61	28.29 ± 7.11	0.17
Height (m)	1.66 ± 0.05	1.68 ± 0.04	0.27
Weight (kg)	68.35 ± 15.23	64.04 ± 4.57	0.24
BMI (kg/m²)	24.66 ± 4.99	22.75 ± 2.03	0.18

*Data presented as mean ± SD*

## Results: Braced vs. Unbraced

- Patients improved by average 16% (82.25 mm²)
- Patients worsened by average 18% (110.29mm²)

	IMPROVED (N= 10)	WORSENE (N = 7)
Braced COPEA (mm²)	426.64 ± 22.71	616.06 ± 75.89
Unbraced COPEA (mm²)	508.89 ± 31.95	505.77 ± 28.18
P	0.002	0.04

*Data presented as mean ± SD*

## Results: Pre-Op Physical Exam

- No differences between improved vs. worsened patients

	IMPROVED (N=10)	WORSENE (N=7)	P
Pain Duration (mo)	16.50 ± 11.11	10.71 ± 5.35	0.22
Hip Flexion (deg)	91.70 ± 13.33	91.43 ± 16.14	0.49
Hip Internal Rotation (deg)	6.50 ± 5.80	10.00 ± 5.77	0.12

*Data presented as mean ± SD*

## Results: Pre-Op Radiographic Indices

- No differences between improved vs. worsened patients

	IMPROVED (N=10)	WORSENE (N=7)	P
Center Edge Angle	35.70 ± 4.83	29.29 ± 9.55	0.08
Lateral Center Edge Angle	28.40 ± 5.30	28.43 ± 5.32	0.50
Acetabular Index	38.80 ± 6.56	42.71 ± 3.35	0.08

*Data presented as mean ± SD*

## Results: Intra-Op Findings

- No differences between improved vs. worsened patients

	IMPROVED (N=10)	WORSENE (N=7)	P
CSI Acetabulum	361.65 ± 236.52	201.14 ± 65.19	0.06
CSI Femoral Head	163.55 ± 146.75	172.00 ± 289.32	0.47
CSI Total	525.20 ± 377.89	373.14 ± 289.70	0.19
IP Release (Y/N)	7 / 3	6 / 1	

Data presented as mean ± SD

## Conclusions

- Some patients benefit from off-the-shelf bracing in the form of balance control
- Certain patients benefit from the protective range of motion function of the brace early on in the rehab cycle, but should come out of the brace when achieving independent ambulation
- Future research needed to define the role of a custom (vs off-the-shelf) hip orthosis for the group that did not show benefit at four weeks after surgery

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## Questions



## **Risk Factors for Infection following Knee Arthroscopy: Analysis of a Large U.S. Cohort**

Carter Clement, Kevin Haddix, Alexander Creighton,  
Jeffrey Spang, Joshua Tennant, Ganesh Kamath



North Carolina  
Orthopaedic Association

Annual Meeting – Kiawah Island, SC  
Nov 2015

## **Background**

- Knee arthroscopy is extremely common
- Infections rare
  - Cited as low as 0.04%
  - But potentially devastating
- Risk factors for infection unknown
  - Historically difficult to study due low incidence

## **Goal**

To identify risk factors for infection  
following knee arthroscopy

## **Methods**

- An administrative healthcare database was used
  - Pearlriver, Fort Wayne, IN, USA
  - Complete records from a large private insurer
  - 5% Medicare sample
- Patients identified by CPT code
  - 20 codes representing knee arthroscopy procedures

## **Methods**

- Patients identified by CPT codes
  - 20 codes representing knee arthroscopy procedures
  - Closed procedures considered low-risk (13)
    - e.g. synovectomy, chondroplasty, microfracture
  - Partially open procedures considered high-risk (7)
    - e.g. mosaicplasty or ACL/PCL

## **Methods**

- Infections developed within 90 days
- Deep infections
  - CPT code for I&D
- Superficial infections
  - ICD-9 infection code without CPT for I&D

## Methods

- Infected vs. non-infected patients compared by:
  - Age
  - Sex
  - Diabetes
  - Overweight/obesity
  - Tobacco use
  - Comorbidities (using Charlson Index)
  - High-risk vs. low-risk procedures

## Results

- 433,423 patients underwent 501,691 knee scopes
- Deep infection rate 0.20%
- Superficial infection rate 0.26%

## Results

Deep Infections		
	Relative Risk	95% CI
High-risk procedures	<b>2.27</b>	<b>(1.98-2.60)</b>

## Results

Charlson Comorbidity Index		
	Median (IQR)	Compared to "No Infection"
No Infection	2 (0,3)	-
Deep Infection	1 (0,2)	P = 0.074
Superficial Infection	1 (0,3)	<b>P &lt; 0.001</b>
All Infections	1 (0,2)	<b>P &lt; 0.001</b>

## Conclusion

- Largest risk factor = "high risk" procedure (RR 2.27)
- Other risks: male sex, tobacco user, diabetes, morbid obesity, CCI, and age under 50 years
  - Age likely confounded by procedure risk
- Helpful for pre-op counseling
- May aid in patient selection
- Can facilitate infection prevention efforts by targeting high-risk patients

**Thank You**



## Appendices

(Manuscript Tables)

## Appendix I

Code	Procedure includes:	Number of Procedures including code (at least once)	Rate (of Arthroscopies)	Relative risk of infection
29866	Mosaicplasty with Autograft	1,048	0.21%	High
29867	Mosaicplasty with Allograft	935	0.19%	High
29868	Meniscal Transplantation	310-319	0.06%	High
29870	Synovial Biopsy	9,477	1.89%	Low
29873	Lateral Release	23,430	4.67%	Low
29874	Removal of Loose Body	22,905	4.57%	Low
29875	Limited Synovectomy	59,269	11.81%	Low
29876	Major Synovectomy	50,191	10.00%	Low
29877	Chondroplasty	154,158	30.73%	Low
29879	Abrasion Arthroplasty (Chondroplasty +/- Microfracture)	53,545	10.67%	Low
29880	Meniscectomy, Med and Lat	117,660	23.45%	Low
29881	Meniscectomy, Med or Lat	294,637	58.73%	Low
29882	Meniscal Repair, Med or Lat	18,554	3.70%	High
29883	Meniscal Repair, Med and Lat	1,968	0.39%	High
29884	Lysis of Adhesions	5,381	1.07%	Low
29885	Drilling & Grafting for OCD	535	0.11%	Low
29886	Drilling for OCD	1,321	0.26%	Low
29887	Drilling for OCD with Internal Fixation	1,376	0.27%	Low
29888	ACL Reconstruction	69,428	13.84%	High
29889	PCL Reconstruction	1,172	0.23%	High

## Appendix II

CPT Codes for Knee Incision & Drainage Procedures	
Code	Description
29871	Arthroscopic Knee I&D
27310	Open Knee Arthrotomy
10180	Complex and/or Postoperative I&D
ICD-9 Codes for Postoperative infection	
Code	Description
711.0	Septic Arthritis
998.51	Postoperative Seroma
998.59	Other Postoperative Infection
999.3	Other Infection due to Medical Care

## Appendix III

Diabetes	
Codes (ICD-9)	Description
250.00 - 250.93	Multiple Diabetic Diagnoses
Overweight/Obesity	
Codes (ICD-9)	Description
278.02	Overweight
V85.21 - V85.25	BMI 25-30 (Overweight)
278.00	Obesity Not otherwise Specified
259.9	Obesity of endocrine origin
V85.30 - V85.39	BMI 30-40 (Obese)
278.01	Morbid Obesity
V85.41 - V85.45	BMI 40+ (Morbidly Obese)
Tobacco Use	
Codes	Description
305.1	ICD-9 Code for tobacco use disorder
V15.82	ICD-9 Code for history of tobacco use
99406	
99407	
99411	CPT Codes for smoking cessation counseling
99412	

## Appendix IV

Number of Patients undergoing arthroscopy	433,423
Number of Arthroscopic Procedures	501,691
Number of arthroscopic codes used	1,682,466
Average number of arthroscopic codes per procedure	3.35
Number undergoing I&D's within 90 days	1001
Rate	0.20%
Number of infections within 90 days not requiring I&D	1310
Rate	0.26%
Total number of infections within 90 days	2311
Rate	0.46%

## Appendix V

Age Group (in years)	ICD-9 Code	Rate	95% CI	Relative Risk	95% CI
15-19	711.0	0.00%	0.00%	0.00%	0.00%
20-24	711.0	0.00%	0.00%	0.00%	0.00%
25-29	711.0	0.00%	0.00%	0.00%	0.00%
30-34	711.0	0.00%	0.00%	0.00%	0.00%
35-39	711.0	0.00%	0.00%	0.00%	0.00%
40-44	711.0	0.00%	0.00%	0.00%	0.00%
45-49	711.0	0.00%	0.00%	0.00%	0.00%
50-54	711.0	0.00%	0.00%	0.00%	0.00%
55-59	711.0	0.00%	0.00%	0.00%	0.00%
60-64	711.0	0.00%	0.00%	0.00%	0.00%
65-69	711.0	0.00%	0.00%	0.00%	0.00%
70-74	711.0	0.00%	0.00%	0.00%	0.00%
75-79	711.0	0.00%	0.00%	0.00%	0.00%
80-84	711.0	0.00%	0.00%	0.00%	0.00%
85-89	711.0	0.00%	0.00%	0.00%	0.00%
90-94	711.0	0.00%	0.00%	0.00%	0.00%
95-99	711.0	0.00%	0.00%	0.00%	0.00%
Sex					
Male					
Female					
Comorbidity					
No comorbidity					
Diabetes					
Obesity					
Tobacco Use					
Postoperative Infection					

## Appendix V (abridged)

Table 5. Post-Arthroscopic Knee Infections by Age and Sex (2005-2012)									
	Deep Infections			Superficial Infections			Total Infections		
	P-Value	Relative Risk	95% CI	P-Value	Relative Risk	95% CI	P-Value	Relative Risk	95% CI
High-risk procedures	<0.001	2.27	(1.98-2.60)	<0.001	2.13	(1.90-2.40)	<0.001	2.19	(2.00-2.39)
Known tobacco use	<0.001	1.68	(1.39-2.03)	<0.001	1.76	(1.50-2.06)	<0.001	1.72	(1.53-1.95)
Sex (Male vs. Female)	<0.001	1.67	(1.47-1.90)	0.031	1.13	(1.01-1.26)	<0.001	1.33	(1.23-1.45)
Overweight	0.003-0.863	0.43-2.38	(0.11-4.30)	0.388	1.36	(0.68-2.71)	0.009-0.884	0.95-1.80	(0.51-2.83)
Obese	0.072-0.572	0.74-0.92	(0.54-1.23)	0.033	1.27	(1.02-1.59)	0.218-0.673	1.04-1.12	(0.87-1.33)
Morbidly Obese	0.001	1.57	(1.19-2.09)	<0.001	1.77	(1.40-2.24)	<0.001	1.68	(1.40-2.01)
Diabetes	<0.001	1.53	(1.29-1.82)	0.002	1.29	(1.10-1.51)	<0.001	1.39	(1.24-1.56)
Age (<50 vs. >50 yrs)	<0.001	1.38-1.43	(1.21-1.63)	<0.001	1.38-1.41	(1.24-1.57)	<0.001	1.38-1.43	(1.27-1.56)

## Appendix VI

Table 6. Charlson Comorbidity Index (CCI) by Infection Status		
Infection Status	CCI Median (IQR)	P-Value (Compared to "No Infection")
No Infection	2 (0,3)	-
Infection Undergoing I&D	1 (0,2)	0.074 <sup>1</sup>
Infection not Undergoing I&D	1 (0,3)	<0.001 <sup>2</sup>
All Infections	1 (0,2)	<0.001 <sup>2</sup>

<sup>1</sup>Trend toward patients with infections having higher CCI despite lower median because data distribution skewed with long upper tail

<sup>2</sup>Patients with infections have higher CCI despite lower median because data distribution skewed with long upper tail

No difference between infections undergoing or not undergoing I&D (P=0.968)



Daryl Henshaw, MD  
Department of Anesthesiology  
Wake Forest University School of Medicine  
October 10<sup>th</sup>, 2015

## Randomized Prospective Study of Anesthetic Techniques in Unicondylar Knee Arthroplasty

## Disclosures

- No conflicts of interest related to the material presented in this presentation.

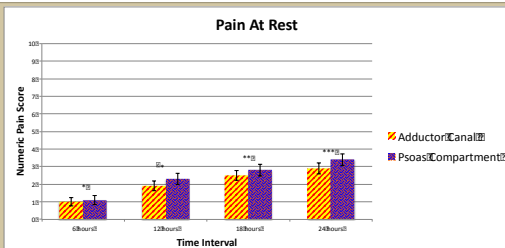
## Study Details

- Prospective, randomized, double-blind equivalency trial
- Medial unicondylar knee arthroplasty
- 150 patients (147 analyzed)
  - 75 Psoas compartment block
  - 75 Adductor canal block
- All had posterior capsule injection
- Multimodal analgesics
  - Celecoxib, Acetaminophen, Pregabalin

## Study Details

- Primary outcome:
  - Pain scores with rest and movement at 6 hours
    - Within 2 points on verbal pain scale (0-10 range)
- Secondary outcomes:
  - Pain scores at 12, 18 & 24 hours (rest & movement)
  - Quadriceps strength (0-5 scale) @ 6 hours
  - Opioid consumption and opioid related side effects over 24 hour period

## Rest Pain

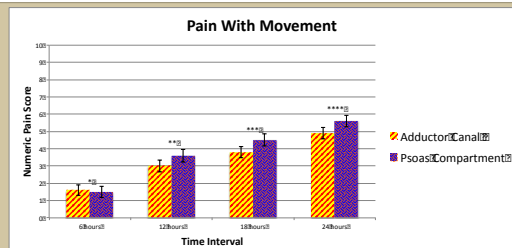


Mean verbal pain scores (Numerical Rating Scale 0-10) at 6-, 12-, 18-, and 24-hours.  
Error bars represent standard error of the mean.

\* $P < 0.0001$ , \*\* $P = 0.0003$ , \*\*\* $P = 0.001$ ;

$P < 0.025$  denotes equivalency at 6 hours;  $P < 0.05$  denotes equivalence at 12, 18 and 24 hours

## Movement Pain



Mean verbal pain scores (Numerical Rating Scale 0-10) at 6-, 12-, 18-, and 24-hours.  
Error bars represent standard error of the mean.

\* $P < 0.0001$ , \*\* $P = 0.0022$ , \*\*\* $P = 0.0045$ , \*\*\*\* $P = 0.0026$

$P < 0.025$  denotes equivalency at 6 hours;  $P < 0.05$  denotes equivalence at 12, 18 and 24 hours

## Quadriceps Strength

	Adductor Canal Block (n=74)		Psoas Compartment Block (n=73)		P-value		
	n	%	n	%			
Straight leg raise (score)*							
5	29	39	8	11	<0.0001*		
4	24	32	7	10	0.001*		
3	13	18	21	29	0.12		
2	6	8	18	25	0.008*		
1	2	3	19	26	<0.0001*		
(5 or 4)	53	72	15	21	<0.0001*		
(3 or 2 or 1)	21	28	58	79	<0.0001*		
	Mean	SD	Median	Mean	SD	Median	
Straight leg raise (score)	4.0	1.1	4.0	2.5	1.3	2	<0.0001*

## Opioids and Side Effects

- No differences in:
  - Time to first analgesic.
  - Cumulative opioids consumed over 24 hours.
  - Incidence of nausea or vomiting at any time point.
  - Incidence of itching at 12,18 or 24 hours.
- Only difference found:
  - Higher incidence of itching at 6 hours in adductor canal group; ( $p=.046$ )

## Conclusion

- Adductor canal blockade:
  - Equivalent analgesia to a psoas compartment block.
  - Significantly less quadriceps motor weakness.
  - Similar side effect profile, except for increased itching at 6 hours.
- Should be considered as an analgesic option for patients undergoing medial unicompartmental arthroplasty.

# MRI findings versus intra-operative pathology in hip arthroscopy

32<sup>nd</sup> Southern Orthopaedic Association Annual Meeting  
Asheville, North Carolina  
July 16, 2015

## Duke Orthopaedic Surgery

Kathleen D Reay, MD  
Julie A Neumann, MD  
Thomas Hash II, MD  
Steven A. Olson, MD



## Disclosures

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## Introduction

- MRIs and MRAs are standard of care to evaluate intra-articular hip pathology
- Arthroscopy: gold standard to evaluate hip labral pathology
- MRI has been reported to accurately identify labral pathology in 91-95% of cases
- Determine accuracy of MRI compared to intra-operative labral and chondral pathology specifically in setting of hip dysplasia



## Methods

- Retrospective review
- PI performed all surgeries
- Indications: CEA < 20, failed non-op mgmt X 6 months, mechanical symptoms
- Combined hip arthroscopy and periacetabular osteotomy (PAO) for treatment of intraarticular pathology and hip dysplasia
- January 1, 2013 to December 31, 2013



## Methods

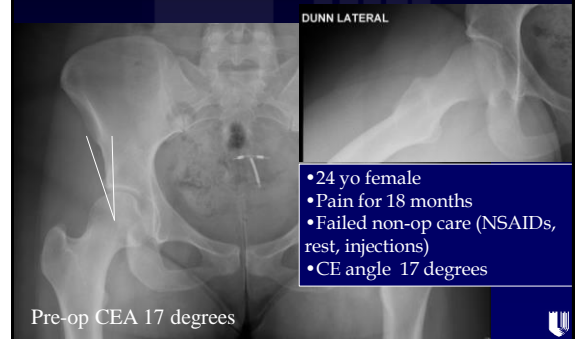
- 17 patients (19 hips)
  - 14 females: 3 males
  - 11 left hips: 8 right hips
- Average age at surgery 29.49 years (range, 17-42 years)
- Fellowship trained musculoskeletal radiologist blinded to intra-op findings reviewed 19 pre-op MRIs (taken avg 83 days before surgery)
  - MRI findings directly compared to operative note findings



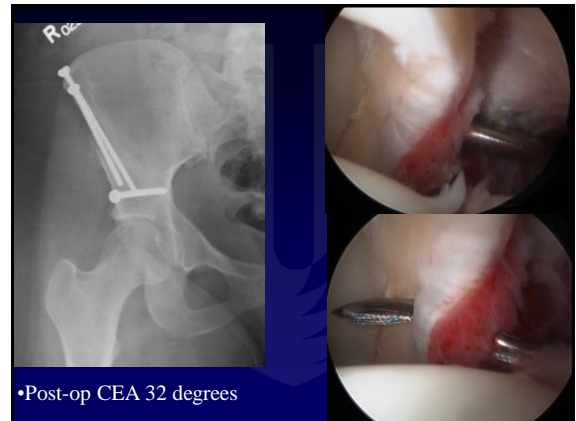
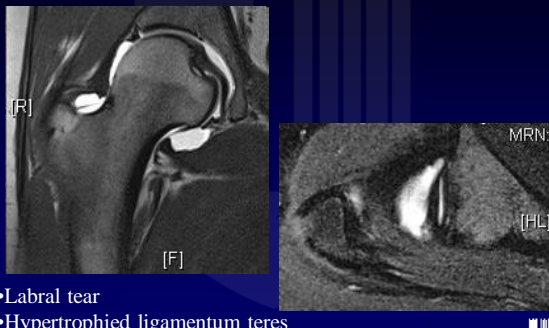
## Operative Procedure

- Supine on fracture table – arthroscopic perineal post
- Single prep and drape – for arthroscopy and PAO
- Hip arthroscopy followed by PAO
- Single surgeon performs both arthroscopy and PAO

## Case Example



## Pre-operative MRI



## Results

- Labrum
  - MRI correctly correlated to intraop pathology 18/19 (PPV = 94.7%) of hips
  - 1 of 19 (5.3%) hips MRI demonstrated labral tear when labrum was intact

## Results

- Chondral lesions
  - Discrepancy between MRI and intra-operative findings in 9/19 (47.4%) of hips
    - MRI noted possible acetabular cartilage delamination vs softening in 4/19 (21.1%) of the hips when intact
    - 3/19 (15.8%) hips MRI showed cartilage delamination with full or partial thickness loss when cartilage was intact
    - 1/19 (5.3%) hips demonstrated questionable acetabular chondral loss when grade III/IV changes were noted in the acetabulum
    - 1/19 (5.3%) hips demonstrated no chondral defect when mild delamination was noted
  - MRI correlated with intraoperative findings in 10/19 (52.6%) hips
    - No chondral defect in the acetabulum in 1/19 (5.3%) of hips when no delamination

## Results

- Ligamentum teres:
  - Discrepancy between MRI and intra-operative findings in 7/19 (36.8%) of hips
    - Frayed, degenerative or partial tear in the ligamentum teres in 5/19 (26.3%) of hips when no tear was noted (hypertrophied or normal)
    - Ligamentum teres rupture in 1/19 (5.3%) hip when a small tear was noted
    - Intact ligamentum teres in 1/19 (5.3%) of hips when hypertrophied
  - Intraoperative findings
    - Torn 4/19 (21.1%)
    - Hypertrophied 8/19 (42.1%)
  - Overall accuracy of diagnosis on MRI was found to be 63.2%

## Overall Results

PPV – 56%  
NPV – 33%  
Sensitivity – 82%  
Specificity – 12.5%  
Accuracy – 52.6%

## Limitations

- Retrospective
- Small cohort
- One MSK radiologist reviewing MRI
- Non-standardized MRI sequences
- Observation, time and selection bias

## Conclusion

- MRI findings compared to intra-operative hip arthroscopy findings
  - Correlate with labral pathology in 94.7% of patients
  - Did not correlate as well for chondral (52.6%) or ligamentum teres (63.2%) pathology
  - Caution when using MRI to diagnose cartilaginous or ligamentum teres pathology
  - Consider having a MSK trained radiologist to review preoperative MRIs

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## Thank You!



### Advantages to Non-arthrographic MRI

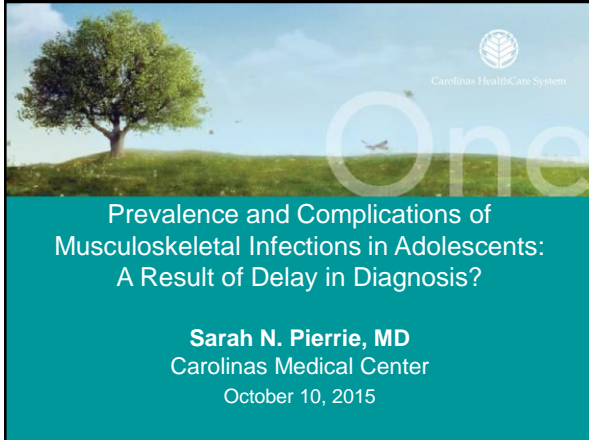
- Noninvasive
- Free of radiation and gadolinium exposure
- Least resource intensive and thus less costly
- Logistically easier to coordinate compared to d-MRA and i-MRA



### HASH MRI Protocol

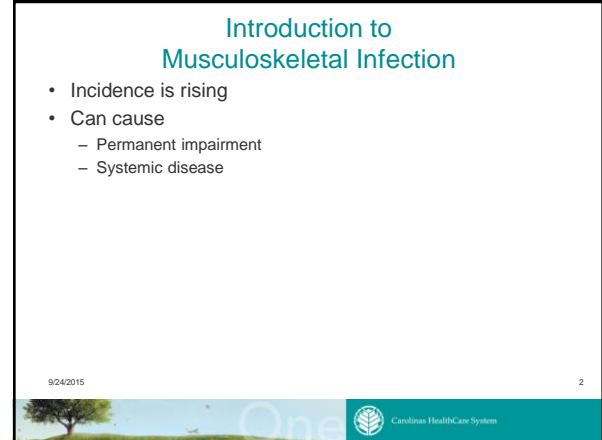
- Cor T1
- Obl Ax fs pd
- Cor fs pd
- Sag fs pd
- Axial VIBE sequence: allows for reformatting
- Ax haste (pd-proton density)





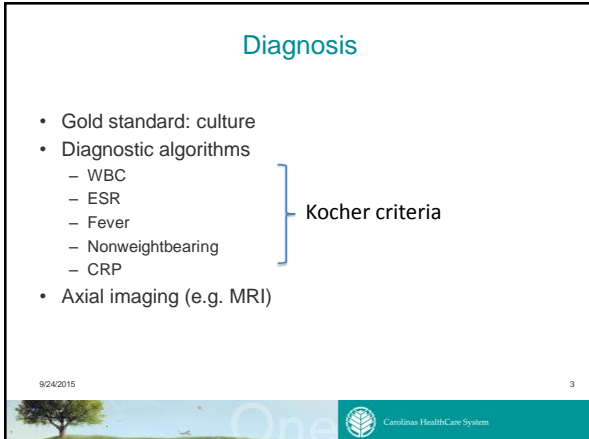
## Prevalence and Complications of Musculoskeletal Infections in Adolescents: A Result of Delay in Diagnosis?

**Sarah N. Pierrie, MD**  
 Carolinas Medical Center  
 October 10, 2015



## Introduction to Musculoskeletal Infection

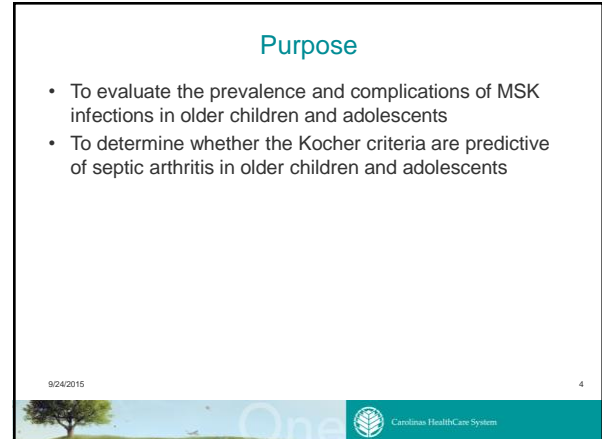
- Incidence is rising
- Can cause
  - Permanent impairment
  - Systemic disease



## Diagnosis

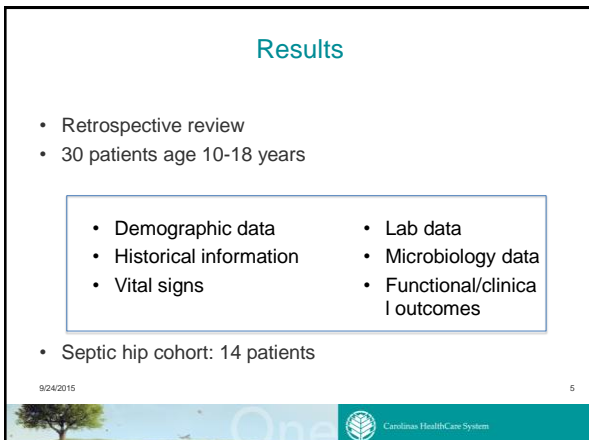
- Gold standard: culture
- Diagnostic algorithms
  - WBC
  - ESR
  - Fever
  - Nonweightbearing
  - CRP
- Axial imaging (e.g. MRI)

} Kocher criteria



## Purpose

- To evaluate the prevalence and complications of MSK infections in older children and adolescents
- To determine whether the Kocher criteria are predictive of septic arthritis in older children and adolescents

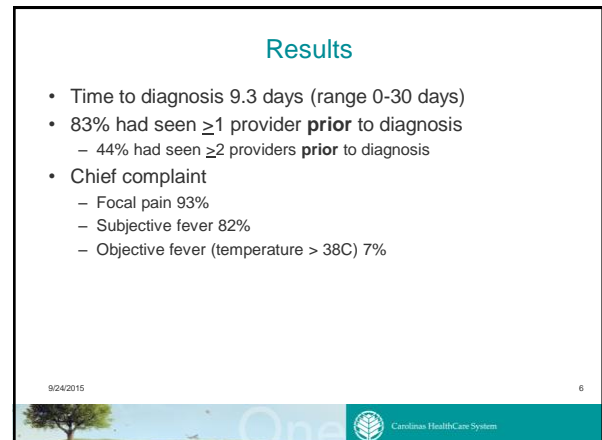


## Results

- Retrospective review
- 30 patients age 10-18 years

• Demographic data	• Lab data
• Historical information	• Microbiology data
• Vital signs	• Functional/clinical outcomes

- Septic hip cohort: 14 patients



## Results

- Time to diagnosis 9.3 days (range 0-30 days)
- 83% had seen  $\geq 1$  provider **prior** to diagnosis
  - 44% had seen  $\geq 2$  providers **prior** to diagnosis
- Chief complaint
  - Focal pain 93%
  - Subjective fever 82%
  - Objective fever (temperature  $> 38^{\circ}\text{C}$ ) 7%



## Results

### Laboratory values

- WBC 11,600
- ESR 50.6
- CRP 15.5

### Cultures

- 68% OSSA
- 9% MRSA

9/24/2015

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## Results

- Patients with...
  - 1 positive criterion: n = 7
  - 2 positive criteria: 6
  - 3 positive criteria: 1
  - 4 positive criteria: 0
- Most common positive finding: ESR
- Mean CRP: 16.8mg/L

9/24/2015

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## Results: Complications

Systemic Complications	
Deep venous thrombosis	3
Pneumonia	2
Pulmonary embolism	1
<i>Clostridium difficile</i> infection	1
Wound infection	1
Musculoskeletal Complications	
Avascular necrosis	3
Persistent limp	3
End-stage arthrosis	2
Heterotopic ossification	2
Loss of terminal motion	2
Recurrent idiopathic effusion	1
Chronic osteomyelitis	1

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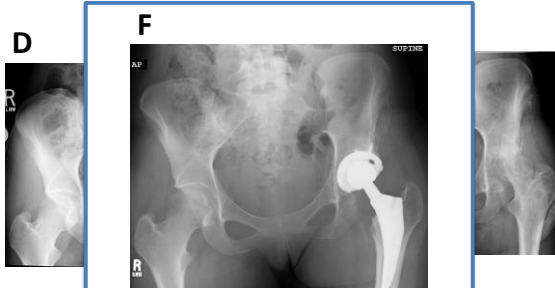
## Complications: AVN



9/24/2015

10

## Complications: AVN



9/24/2015

11

## Discussion

- In teens and adolescents with MSK infection,
- There is a trend toward longer symptom duration before definitive treatment
  - Kocher criteria are inconsistently present
  - CRP is almost uniformly elevated
  - MRI is an important diagnostic tool
  - Complications occur in one third of patients

9/24/2015

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9/24/2015

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Carolina's HealthCare System  
Levine Children's Hospital

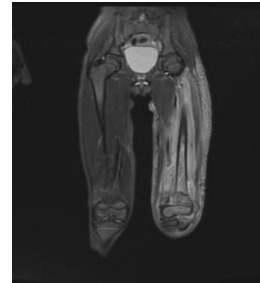
## The Utility of a Screening MRI for Pediatric Patients with Suspected Musculoskeletal Infection

Paulvalery Roulette, MD  
Franklin Gettys, MD,  
Brian Scannell, MD  
Steven Frick, MD  
Nigel Rozario, BS  
Kelly VanderHave, MD  
Brian Brighton, MD, MPH

One

### Background

- Septic Arthritis or Benign Process?
- Multifocal Musculoskeletal Infections
- Utility of MRI



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Levine Children's Hospital

### Study Purpose

- Describe our imaging protocol
- Report on a cohort of patients who underwent a screening MRI for suspected musculoskeletal infection prior to any procedural intervention.
- Determine the clinical and/or laboratory values that are predictive of patients presenting with septic arthritis versus multifocal musculoskeletal infection

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Levine Children's Hospital

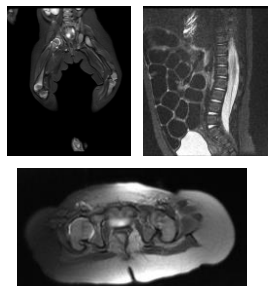
### Methods

- Single institution retrospective review from 2008-2014
- Patients age <19 years
- Include all who underwent a screening MRI for a suspected musculoskeletal infection prior to intervention
  - A screening MRI was defined as an MRI that examined 1 joint and at least 1 continuous bony structure
- Exclusion criteria:
  - Incomplete medical records
  - Incomplete MRI images
  - MRI for non infectious work up

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### MRI Protocol

- Hip to toe
- SAG IR Lumbar
- COR IR, COR T1 Pelvis to Ankles
- AX T1, AX T2 FS
- Gadolinium per MD order or Rad request



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### Analysis

#### Patients were analyzed in 4 groups:

- no infection
- musculoskeletal infection
- septic arthritis of a joint without concomitant infection (SAJ)
- multifocal musculoskeletal infection (MMI)

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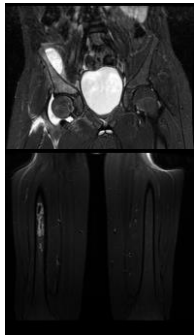
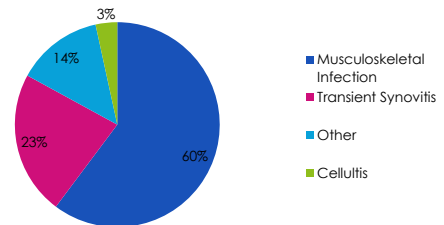
## Analysis

### Independent Variables

- Age, gender
- Weight bearing status (WBS) at admission,
- White blood cell count (WBC), CRP (mg/dL), ESR, hemoglobin (HGB), temperature at admission.
- Number of positive Kocher criteria (temperature > 101.3 F°, ESR > 40 mm/hr, non weight bearing on affected side, WBC ≥ 12,000 cells/mm<sup>3</sup>)

## Results

Study Population n = 88



## Results

### Musculoskeletal Infections

- SAJ 19/53 (35.8%)
  - 13 hip, 4 knee, 1 ankle, 1 tarsal/metatarsal
- MMI 13/53 (24.5%)
  - 12 septic arthritis of hip with
    - osteomyelitis (8),
    - pyomyositis (2)
    - abscess (1)
  - septic arthritis of knee, abscess, osteomyelitis, (1)
  - 1 septic arthritis of knee with osteomyelitis

### Patients with Multifocal Musculoskeletal Infection (MMI) and their Subsequent Procedure

Patient	Procedure
Patient #1	I&D Hip, Abscess, Drilling Femoral Head
Patient #2	I&D Hip, Knee, Ankle, Foot
Patient #3	I&D Hip & Pelvis
Patient #4	I&D Hip & Drilling Ilium
Patient #5	I&D Hip & Bilateral Tibia Aspiration
Patient #6	Hip and Proximal Femur Aspiration
Patient #7	I&D Gluteus Maximus, Minimus, Iliac wing
Patient #8	I&D Hip
Patient #9	I&D Hip
Patient #10	I&D Hip
Patient #11	I&D Hip
Patient #12	I&D Hip
Patient #13	I&D Knee

## Results

### MMI vs. SAJ

- MMI older
  - 7 years MMI vs. 3.3 years SAJ (p=0.09)
- MMI higher CRP
  - 13.09(6.24-19.94) vs.. 4.52(2.83-6.2) p=0.01.
- MMI lower WBC count
  - 12.05(7.33-16.77) vs.. 13.77(12.08-15.45) p=0.02.
- No differences in age, ESR, number of Kocher criteria, and temperature on admission.

## Results

### Those with CRP of 9 were 9.7 times more likely to have MMI

- 14.7 times more likely with CRP of 13
- 22 times more likely with CRP of 15

## Results

### Musculoskeletal infection (MI) vs. No infection (NI)

- CRP
  - (NI) 3.7(1.57-5.83) vs. (MI) 9.66(7.21-12.1)  $p < 0.001$
- ESR
  - (NI) 25.96(18.39-33.53) vs. (MI) 60.36(49.01-71.71)  $p < 0.001$
- Temperature on admission
  - (NI) 98.86(98.06-99.66) vs. (MI) 99.63(99.14-100.13)  $p = 0.02$
- Number of Kocher criteria
  - (NI) 1.14(0.85-1.43) vs. (MI) 1.87(1.56-2.17)  $p = 0.002$



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Lawrence Ch. Identi's Hospital

## Study Limitations

- Retrospective study that only included patients from an MRI database



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## Conclusions

- Screening MRI was useful in the diagnosis and management of patients presenting with musculoskeletal infection.
- Nearly 25% of our cohort of musculoskeletal infections had a multifocal musculoskeletal infection.
- The odds of having a multifocal musculoskeletal infection on MRI versus an isolated septic arthritis was 9.7 times higher with a CRP > 11, however no other clinical factors were significant in determining the presence of an associated bone or soft tissue infection on MRI.
- The use of MRI is recommended in the evaluation and management of children with musculoskeletal infections as clinical factors alone may not be adequate in determining the presence of multifocal infection.
- The early recognition of a multifocal infection allows one to make the appropriate diagnosis and provide proper surgical care at the initial operation.



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## Treatment of Pelvic Chondroblastoma with Denosumab: The Role of RANK Signaling in Benign-Aggressive Tumors

Mitchell Klement, MD,  
Julia Visgauss, MD, Will Eward DVM, MD  
NCOA Annual Meeting  
Oct 10, 2015

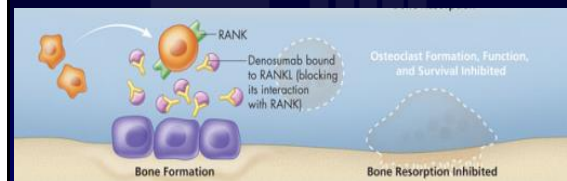
### Case Presentation

- 15 yoM with eight months of progressive, mechanical right hip pain
- PE: antalgic gait and tenderness to palpation over the right buttock and groin

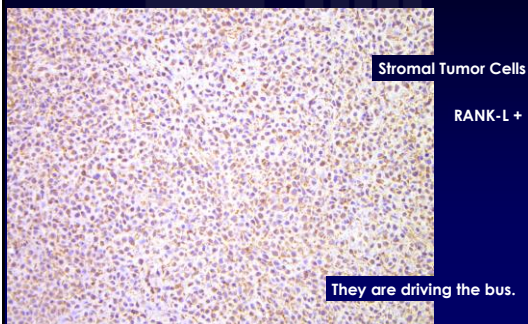
### Case Presentation



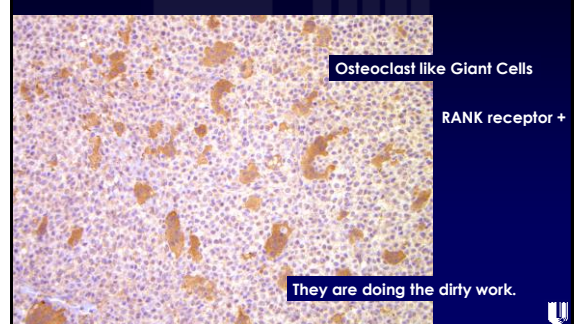
### Rank-Rank L Signaling Pathway



### In Benign Aggressive Tumors, who are the main cellular players?



### In Benign Aggressive Tumors, who are the main cellular players?



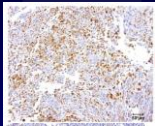


## Role of Denosumab in Benign Aggressive Tumors

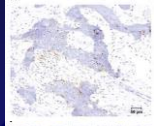
**Denosumab Induces Tumor Reduction and Bone Formation in Patients with Giant-Cell Tumor of Bone**

Daniel G. Branstetter, Scott D. Nelson, J. Carlos Manivel, et al.  
*Clin Cancer Res* 2012;18:4415-4424.

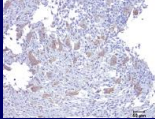
Pre-treatment:  
numerous  
RANK-L positive  
tumor stromal  
cells



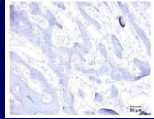
Post-treatment:  
minimal RANK-L  
positive tumor  
stromal cells



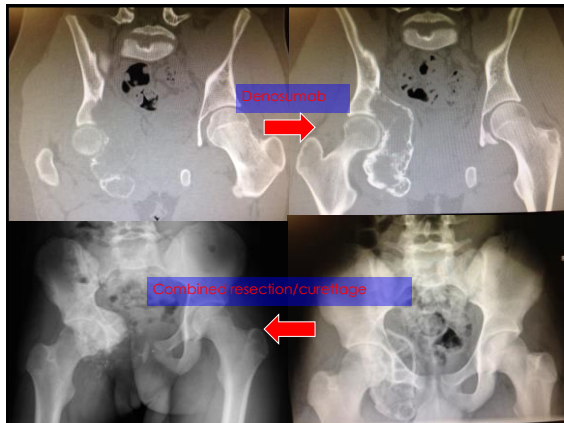
Pre-treatment:  
numerous  
RANK receptor  
positive tumor  
giant cells



Post-treatment:  
No RANK  
receptor  
positive tumor  
giant cells

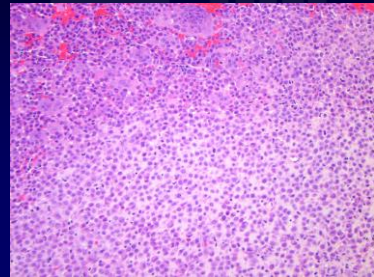


## Back to Our Case Report



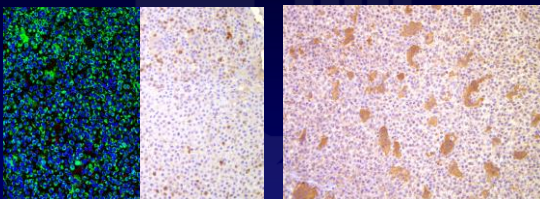
## What is the effect of Denosumab on Chondroblastoma?

Pre-Denosumab Sample



## What is the effect of Denosumab on Chondroblastoma?

Pre-Denosumab Sample

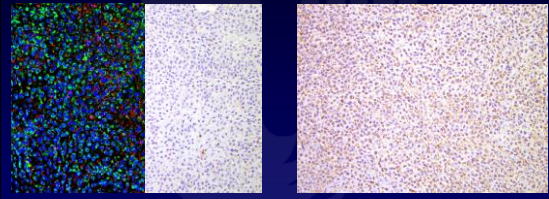


Immunofluorescence and  
immunohistochemical staining  
of RANK-L

Immunohistochemical staining  
of RANK receptor

## What is the effect of Denosumab on Chondroblastoma?

Post-Denosumab Sample



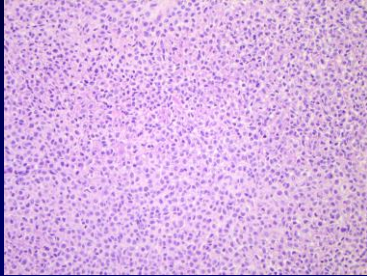
Immunofluorescence and  
immunohistochemical staining  
of RANK-L

Immunohistochemical staining  
of RANK receptor



## What is the effect of Denosumab on Chondroblastoma?

Post-Denosumab Sample



## What is the effect of Denosumab on Chondroblastoma?

- Treatment of Chondroblastoma with Denosumab results in abrogation of osteoclast-like giant cell formation with decrease in resultant osteolysis
- However, UNLIKE GCT, we don't see over-expression of RANK-L in the stromal cells of Chondroblastoma, and see little effect on neoplastic mononuclear cells following treatment with Denosumab.

## Summary

- The RANK/RANK-L signaling pathway plays an important role in the osteolytic process of benign aggressive tumors such as GCT and Chondroblastoma.
- Indications, dosing, and duration of Denosumab treatment are still being investigated
- The success with Denosumab treatment in our patient are encouraging regarding the potential for improvements in treatment and management of patient's with osteolytic pathologic lesions beyond GCT of Bone
- However it's pathologic role in various tumors may be different, which may have implications on future management.

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13. <http://www.headtoplexus.net/article/bone-biology-and-role-rank-ranklig-pathway>

## Acknowledgements

- Brendan Dickson (Univ. Toronto)
- Alex Lazarides (Duke)
- Suzanne Bartholf (Duke)
- Jason Somarelli (Duke)

**Risk Factors For Disease Progression After Surgical Treatment of Extremity Metastatic Bone Disease**

Elizabeth Scott, Mitchell R. Klement, MD, Brian E. Brigman, MD, PHD, William C. Eward, MD DVM

Elizabeth Scott, BA - noting to disclose  
 Mitchell R. Klement, MD - nothing to disclose  
 Brian E. Brigman, MD  
 Lumicell Diagnostics: Research support  
 Musculoskeletal Transplant Foundation: Paid consultant; Paid presenter or speaker; Research support  
 Musculoskeletal Tumor Society: Board or committee member  
 Plexicon: Paid consultant  
 William C. Eward, MD DVM - nothing to disclose

**Purpose / Hypothesis**

one-year postoperative survival for metastatic skeletal disease has improved with better adjuvant therapies:

0.3% Wedin, R., et al 1999 → 36% Ratasvuori et al 2013

We aimed to assess risk factors associated with:

- radiographic disease progression
- surgical failure
- survival

in "longer-term" (6+ month) survivors


**Materials and Methods**

**Retrospective Review** surgical treatment of 89 metastatic bone lesions, extremity only treated between 2004-2014 by Duke Orthopaedic Oncology 6+ months of radiographic imaging

**Variables Considered** lesion characteristics, surgical details, radiotherapy timing, antiresorptive medication use, Mirels score

**Statistical Analysis** Fischer's Exact Test, t-test, Cox proportional-hazards models

- Radiographic Disease Progression**  
 plain radiographs assessed at 3 month intervals  
 Harada, H., et al. (2010) criteria
- Surgical Failure**  
 hardware failure, infection
- Survival**



Interval Disease Progression & Implant Failure 18 months postop

**Results**

**1. Radiographic Disease Progression**

- Gender (Female): HR 0.361, p<0.01
- Round Cell Cancer (vs. RCC): HR 0.441
- Non-Renal Carcinomas (vs. RCC): HR 0.481
- Humeral Lesions (vs. Femoral): HR 0.399
- Bisphosphonates (vs. Denosumab): HR 0.422

**2. Surgical Failure (Fisher)**

- Tumor Origin p=0.005
- Extremity p=0.059
- Gender p=0.087
- Procedure p=0.066
- Mirels Score p=0.048

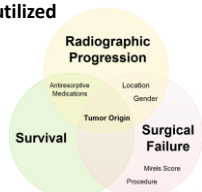
**3. Survival**

- Antiresorptive Use (BP) HR 0.504, p=0.020
- Tumor Origin p=0.05
  - Lung (vs. RCC) HR 3.32, p=0.025
  - Prostate (vs. RCC) HR 3.22, p=0.066

Failure & Radiographic Progression Rates		
	Failure	Progression
<b>Overall</b>	<b>16.85%</b>	<b>43.82%</b>
<b>Renal Cell Carcinoma</b>	30.77%	61.54%
<b>Other Carcinomas:</b> Breast, Prostate, Thyroid, Lung	12.0%	36.0%
<b>Round Cell Cancers:</b> Multiple Myeloma & Lymphoma	3.03%	30.3%
<b>Male</b>	25.71%	68.57%
<b>Female</b>	11.11%	27.78%
<b>Femur</b>	19.35%	48.39%
<b>Humerus</b>	0%	21.50%
<b>All Fixation</b>	22.22%	48.1%
<b>IM Nail</b>	24.32%	48.6%
<b>Endoprosthesis</b>	8.57%	37.1%
<b>Denosumab Use</b>	11.54%	60.0%
<b>Bisphosphonate Use</b>	20.0%	30.7%

**Conclusion**

- Endoprosthetic Replacement > Fixation for Some Groups**
  - **Tumor Type:** Renal Cell Carcinoma > Non-Renal Carcinomas > Round-Cell
  - **Location:** Femoral > Humeral lesions
  - **Gender:** Men > Women
- Antiresorptive Therapy should be utilized**
  - strong association with **patient survival**
  - increased time to **radiographic progression** (bisphosphonates > denosumab)



**Questions?**

Elizabeth Scott, BA  
[elizabeth.j.scott@dm.duke.edu](mailto:elizabeth.j.scott@dm.duke.edu)

Dr. William Eward, MD, DVM  
[will.eward@dm.duke.edu](mailto:will.eward@dm.duke.edu)



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