

# Walter Reed Bethesda ORTHOPAEDIC Journal



2013

Walter Reed  
Bethesda  
Orthopaedic  
Alumni Association

**Contents**

**2013**

President's Letter . . . . .	iii
Chairman's Letter . . . . .	iv
Residency Program Director's Letter . . . . .	v
Residents/Fellows . . . . .	vi
2013 Active Staff. . . . .	vii
Editorial. . . . .	viii

**CHIEF RESIDENT PAPERS**

**Subpectoral Biceps Tenodesis: an Anatomical Study and Evaluation of At-Risk**

<b>Structures. . . . .</b>	<b>1</b>
----------------------------	----------

MAJ Jonathan F. Dickens, MD, MC, USA; MAJ Kelly G. Kilcoyne, MD, MC, USA;  
LCDR Scott M. Tintle, MD, MC, USN; MAJ Jeffrey Giuliani, MD, MC, USA;  
COL Richard A. Schaefer, MD, MC, USA; CDR John-Paul Rue, MD, MC, USN

**Complications and Reoperations Associated with Combat-Related Tibial Plafond**

<b>Fractures . . . . .</b>	<b>5</b>
----------------------------	----------

CPT Husain Bharmal, MD; LT Louis Lewandowski, MD; MAJ Korboi N. Evans, MD, MS;  
LCDR Steven Grijalva, MD; CDR Mark E. Fleming, DO, MC, USN

**Reported Concussion Rates for Three Division I Football Programs: an Evaluation  
of New NCAA Concussion Policy . . . . .**

<b>10</b>
-----------

MAJ Kelly G. Kilcoyne, MD; MAJ Jonathan F. Dickens, MD; LTC Steven J. Svoboda, MD;  
LTC Brett D. Owens, MD; LTC Robert T. Sullivan, MD; CDR John-Paul Rue, MD

**BMP Regulation of Mesenchymal Progenitor Cells in Regenerating Muscle Tissue . . . . .**

<b>12</b>
-----------

CPT Matthew W. Kluk, MD, MC, USA; Youngmi Ji, PhD; Orna Amrani, MD;  
Wesly M. Jackson, PhD; LTC Leon J. Nesti, MD, PhD, MC, USA

**Application of the Orthoplastic Reconstructive Ladder to Preserve Lower Extremity**

<b>Amputation Length. . . . .</b>	<b>15</b>
-----------------------------------	-----------

CDR Mark E. Fleming, MD, MC, USN; LCDR J. Anthony O'Daniel, MD, MC, USN;  
CPT Husain Bharmal, MD, MC, USA; LCDR Ian Valerio, MD, MC, USN

**Heterotopic Ossification Resection after Open Peri-articular Combat-Related**

<b>Elbow Fractures . . . . .</b>	<b>19</b>
----------------------------------	-----------

LCDR(s) Kevin W. Wilson, MD, MC, USN; MAJ Jonathan F. Dickens, MD, MC;  
LCDR Reed Heckert, MD, MC, USN; LCDR Scott M. Tintle, MD, MC, USN;  
CDR(r) John J. Keeling, MD, MC, USN; COL Romney C. Andersen, MD, MC, USN;  
MAJ(P) Benjamin K. Potter, MD, MC, USA

## RESEARCH FELLOW PAPERS

<b>An Innovative Culture System for the Study of Post-Traumatic Heterotopic Ossification . . . . .</b>	<b>23</b>
CPT Husain M. Bharmal, MD, MC, USA; Gregory T. Christopherson, PhD; Youngmi Ji, PhD; Wesley M. Jackson, PhD; LTC Leon J. Nesti, MD, PhD	
<b>Outcomes Following Cervical Disc Arthroplasty: a Retrospective Review . . . . .</b>	<b>26</b>
CPT John P. Cody, MD; LCDR Robert W. Tracey, MD; CPT Daniel G. Kang, MD; CPT Adam J. Bevevino, MD; LTC Michael K. Rosner, MD; LTC Ronald A. Lehman, Jr., MD	
<b>The Use of Dilute Dakin's Solution for the Treatment of Invasive Fungal Infection in the Combat Wounded: a Case Series . . . . .</b>	<b>29</b>
LT Louis Lewandowski, MD, MC, USN; CPT Richard Purcell, MD, MC, USA; CDR Mark Fleming, DO, MC, USN; Lt Col Wade T. Gordon, MD, MC, USAF	
<b>Neurovascular Entrapment due to Combat-Related Heterotopic Ossification. . . . .</b>	<b>31</b>
CPT Elizabeth M. Polfer, MD; CDR Jonathan A. Forsberg, MD; CDR Mark E. Fleming, DO; MAJ(P) Benjamin K. Potter, MD	
<b>Outcomes of Single-Level Cervical Disc Arthroplasty versus Anterior Cervical Discectomy and Fusion: a Single Center, Retrospective Review . . . . .</b>	<b>37</b>
LCDR Robert W. Tracey, MD; CPT John P. Cody, MD; CPT Daniel G. Kang, MD; CPT Adam J. Bevevino, MD; LTC Michael K. Rosner, MD; LTC Ronald A. Lehman, Jr., MD	

## BEST PAPERS

<b>Cervical Posterior Foraminotomy's Effect on Segmental Range of Motion in the Setting of Total Disc Arthroplasty . . . . .</b>	<b>41</b>
CPT Adam J. Bevevino, MD; LTC Ronald A. Lehman, Jr., MD; CPT Daniel G. Kang, MD; Divya Ambati, MS; Rachel Gaume, BS; CDR David E. Gwinn, MD; Anton E. Dmitriev, PhD	
<b>Risk Factors for Infection and Amputation following Open, Combat-Related Calcaneus Fractures . . . . .</b>	<b>45</b>
MAJ Jonathan F. Dickens, MD, MC, USA; MAJ Kelly G. Kilcoyne, MD, MC, USA; CPT Matthew W. Kluk, MD, MC, USA; Lt Col Wade Gordon, MD, MC, USAF; LTC(P) Scott B. Shawen, MD, MC, USA; MAJ(P) Benjamin K. Potter, MD, MC, USA	
<b>The Ventral Lamina and Superior Facet Rule: a Morphometric Analysis for Ideal Thoracic Pedicle Screw Starting Point . . . . .</b>	<b>50</b>
CPT Daniel G. Kang, MD; LTC Ronald A. Lehman, Jr., MD; Lawrence G. Lenke, MD; Rachel Gaume, BS; MAJ Haines Paik, MD	

<p>The views expressed in this journal are those of the authors and do not necessarily reflect the official policy or position of the Departments of the Navy, Army, or Air Force, the Department of Defense, nor the U.S. Government.</p>
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# President's Letter

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## Walter Reed Bethesda Orthopaedic Alumni Association

Dear Members of the Walter Reed Bethesda Orthopedic Alumni Association,

Although long overdue, we now have an Alumni Association. This is a fitting culmination of an overwhelmingly successful merger of the Walter Reed and Bethesda orthopaedic residency training programs. To those who have worked hard to establish this organization, I extend my sincerest thanks. It is truly an honor to serve as your inaugural president.

As a training program, I am convinced that our best work is ahead of us. At this time, with our country still engaged in war, our training program finds itself with the opportunity to reaffirm its mission – a mission that ultimately focuses on developing a military orthopedic surgeon capable of operating in and withstanding the demands of leading a surgical unit in combat. We can now visualize the nature and the complexities of delivering orthopaedic surgical combat care in the 21<sup>st</sup> century and can therefore work backward from there to the operating rooms, clinics, and labs of Walter Reed National Military Medical Center. We have an understandable context for clearly defining the vision of our training program and the Department of Orthopaedic Surgery. From there, we have an exceptional and unremitting responsibility to export this common vision and leadership through our graduates to all of our military hospitals and medical centers as well as the battlefield, creating a collaborative effort and unity of purpose. This will provide an opportunity to examine and refine our research priorities and processes with a principled and disciplined approach. This is the chance to truly make a difference – to establish an enduring legacy.

I look forward to seeing all of you at what promises to be an exceptional first annual meeting in September.

With gratitude and warmest wishes,

A handwritten signature in black ink, appearing to read "Martin Deafenbaugh".

Martin Deafenbaugh, MD

CAPT, MC, USN (ret)

President, Walter Reed Bethesda Orthopaedic Alumni Association



# Chairman's Letter



## DEPARTMENT OF ORTHOPAEDICS

Walter Reed National Military Medical Center  
Bethesda, MD

Alumni of Walter Reed Bethesda Orthopaedics,

Many thanks for your interest in Walter Reed Bethesda Orthopaedics. Over the past year there have been many changes and events in orthopaedics.

First and foremost, the Department of Orthopaedics was established. This was an evolution that occurred over a number of years from the Army perspective, and back to the norm from the Navy perspective. From the mid 1900s to 2001, the Army had an Orthopaedic Surgery Service, while the Navy was a separate department. In 2001, the Army established a Department of Orthopaedics and Rehabilitation, and when our two facilities integrated in 2011, the model of a Department of Orthopaedics and Rehabilitation was chosen for our present institution. Over the years, the size of the department grew enormously, not only due to the current war, but also through the expansion of rehabilitation services offered. In late 2012, the structure of the department was evaluated by the command and it was felt that the Department of Orthopaedics and Rehabilitation would be better served by two separate departments. On January 1, 2013, the Department of Orthopaedics and the Department of Rehabilitation were established. With this initiation, seven services were developed within the Department of Orthopaedics, representing the major areas of orthopaedic sub-specialization.

This year also saw the initiation of the Walter Reed Bethesda Orthopaedic Alumni Association, of which this journal is a product. The mission of the association is to promote camaraderie of past and present members of Walter Reed Bethesda Orthopaedics, have an annual retreat with continuing medical education, publish an annual journal, and promote education. Membership is open to anyone who has served within the former National Naval Medical Center Department of Orthopaedics, the Walter Reed Army Medical Center Orthopaedic Surgery Service, the Walter Reed National Military Medical Center Orthopaedic Surgery Service, or the newly formed Walter Reed Department of Orthopaedics. Members can be residents, staff physicians, physician assistants, nurses, enlisted staff, or support staff. Service is defined as anyone who worked within our organization, including physicians who have been credentialed here while on reserve duty or who assisted with casualty care.

With regard to the things that make an academic orthopaedic department excel (patient care, education, and research), Walter Reed Orthopaedics has certainly flourished. Over the past year, our staff has cared for thousands of patients and been the primary facility treating our country's wounded warriors. Our residents continue to score in the top 25% of all orthopaedic training programs. With regard to research and academics, our department outperformed all others in the hospital, with over 100 peer-reviewed publications in the highest level orthopaedic journals, and hundreds of presentations given at national and international meetings.

Romney C. Andersen, M.D.  
COL, MC, USA

Chairman, Orthopaedic Surgery, Walter Reed National Military Medical Center  
Professor of Surgery, Uniformed Services University  
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<b>COL Romney C. Andersen, MD, USA</b> Chairman, Orthopaedic Surgery
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LCDR James H. Flint, MD
CPT Daniel G. Kang, MD
LCDR Reed M. Heckert, MD
CPT Emily H. Shin, MD

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# Residency Program Director's Letter

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*Residency Program Director*  
**COL SCOTT B. SHAWEN**

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*Chief, Orthopaedic Surgery*  
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**CPT B. SMITH**

**LCDR T. STEELMAN**

**NATIONAL CAPITAL CONSORTIUM  
ORTHOPAEDIC SURGERY RESIDENCY  
Walter Reed National Military Medical Center  
8901 Wisconsin Avenue, Bethesda, MD 20889-5000  
Office (301) 295-8588 Fax (301) 319-2699**

The orthopaedic surgery residency program at the Walter Reed National Military Medical Center has never been stronger. Over the last four years, numerous changes have happened to our immediate hospital system, combining Walter Reed Army Medical Center and National Naval Medical Center into the current hospital complex on the Bethesda campus. As a residency program, working with the Department and at that time, Service leadership, we were able to make many changes that have greatly benefitted resident training. Increased PA support, a dedicated trauma team, a revised academic program, broad educational opportunities, and staff physicians dedicated to resident education make this into what I consider to be the best training program in the military.

We are one of the largest residency programs in the military, with three Army and three Navy residents at each training level. The incoming interns for the 2013-14 academic year comprise what I consider to be one of the strongest classes in years. We welcome CPT Christopher Daniels (Army), LT Kyle Nappo (Navy), CPT Rick Purcell (Army), LCDR Jonathan Seavey (Navy), LT Benjamin Wheatley (Navy), and CPT Jared Wolfe (Army) to the Orthopaedic Surgery Team. I feel that each of these individuals is extremely talented and will add tremendously to the residency program as well as the field of orthopaedic surgery in the years to come.

The graduating class of 2013 will be sorely missed as they head out into fellowship and practice. Their efforts made this a very successful year, with an excellent academic program as well as leadership. MAJ Jonathan Dickens is in a sports medicine fellowship at West Point; LCDR Steve Grijalva has transitioned to practice in Pensacola, Florida; MAJ Kelly Kilcoyne is in a shoulder/elbow fellowship at Johns Hopkins; MAJ Matthew Kluk has transitioned to practice at Ft. Hood, Texas; LCDR Anthony O'Daniel has transitioned to practice at Naval Station Great Lakes; and LCDR Kevin Wilson is at Camp Lejeune, North Carolina.

We have had another very successful academic year, with more than 100 publications involving both residents and staff. The basic science labs, including Walter Reed Spine/Biomechanics, National Institutes of Health/NIAMS, and Naval Medical Research Center at Forest Glen as well as the clinical research programs at the Naval Academy and Walter Reed NMMC, continue to be producers of cutting edge research. The residents have received multiple local and national awards for their efforts.

Most recently, Lt Col Wade Gordon has been selected to become the new Program Director for the residency program starting July 1, 2013. He is a great leader with boundless energy, who will lead this program into the future. I look forward to working with the staff and residents for many more years to come.

**COL Scott B. Shawen**

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# Residents/Fellows

To view the 2013 Graduation pictures, go to [www.wrboaa.org](http://www.wrboaa.org).

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## 2013 Graduate Class



**Left to Right:** MAJ Jonathan Dickens, MAJ Matthew Kluk, LCDR Steven Grijalva, MAJ Kelly Kilcoyne, LCDR Joseph Anthony O'Daniel, LCDR(S) Kevin Wilson

## 2013-2014 Chief Resident Class



**Left to Right:** LT James Flint, CPT Emily Shin, CPT Adam Bevevino, CPT Daniel Kang, LCDR Reed Heckert, not pictured: LCDR Keith Alfieri

## 2013-2014 Senior Resident Class



**First Row:** CPT Elizabeth Polfer

**Second Row:** LT Louis Lewandowski, LCDR Robert Tracey, CPT John Cody, CPT Husain Bharmal

## 2013-2014 Research Fellows



**Left to Right:** CPT Donald Hope, CPT Adam Pickett, LT Scott Wagner, LT Benjamin Chi, LT Patrick Jones, CPT Gregory Van Blarcum

## 2013-2014 Junior Resident Class



**Left to Right:** CPT Alaina Brelin, CPT Brett Smith, LT Daniel Griffin, CPT Michael Donohue, LCDR Theodore Steelman, LT Theodora Dworak

# 2013 Active Staff

## Orthopaedic Arthroplasty & Oncology



**Front Row:** Ellyce Johnson, PA; Betty Spacciapoli, RN; Susan Foster, RN  
**Back Row:** CDR M.T. Newman, MD (Chief); CDR Jonathan A. Forsberg, MD; LT COL Kyle Potter, MD; MAJ Andrew Mack, MD

## Hand Surgery



**Left to Right:** MAJ Justin Mitchell, LTC Christina Cawley, LTC Leon Nesti, CDR George Nanos, CAPT Patricia McKay, LTC Derek Ipsen (Chief), and PA-C Merri-Beth Cully

## Orthopaedic Trauma



**Front Row:** LTC Christina Cawley, Lt Col Wade Gordon (Chief), RN Susan Foster  
**Second Row:** CDR Mark Fleming, CDR Robert Beer, MAJ(P) B. Kyle Potter, COL Romney Andersen  
**Third Row:** LCDR Billy Burk, MAJ Jean-Claude D'Alleyrand, CDR Jonathan Forsberg

## Pediatrics/Spine Surgery



**Left to Right:** Morayma Rivera, LCDR Terrance Anderson, MAJ Melvin Helgeson, Barbara Goudarzi, LTC Ronald Lehman, not pictured: LTC Jefferson Jex

## Podiatry



**Front Row:** RN Gloria Martin-Smith, CDR Monique Gourdine-Shaw (Chief)  
**Second Row:** LPN Nancy Soto, LTC Tobias Glister  
**Back Row:** MAJ Bill Michie

## Sports/Shoulder



**Left to Right:** CAPT Dan Hebert, LCDR Michael McCabe, Mrs. Joyce Yesupria, PA Susan Sinnott, PA Kimberly Farnsworth, Mrs. Bridgett Hott, MAJ Jeffrey Giuliani, Col Jeffrey Davila

## Physician Assistants



**Front Row:** Ellyce Johnson  
**Second Row:** Merri-Beth Cully, Susan Sinott  
**Third Row:** LCDR Billy Burk, Kimberly Farnsworth, Thomas Koontz



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# Editorial

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When COL Andersen approached me about starting this journal, I had no idea what it would involve, given our department does not have a history of putting together anything like it. I have to admit I was a little skeptical at first. However, I realized that because of the great things being done here, creating a journal worthy of publication wouldn't be a difficult job. (Unfortunately, there are still a few residents who seem to enjoy other aspects of their job, i.e., patient care, more than they do research.) The professional staff at Data Trace has also made this process a seamless and enjoyable one.

The goals of the journal are:

- 1) Increase awareness of the significant research being performed in the Department of Orthopaedics,
- 2) Provide a brief overview of the department's productivity, and
- 3) Update all alumni and personnel associated with the department on the great things we are doing.

The graduating chief residents and research fellows were instructed to compose a summarized version of the best product they produced during either their entire residency or their research year, respectively. Additionally, three papers were selected from our department's performance at the Society of Military Orthopaedic Surgeons Annual Meeting this past year, representing some of the best our residents have to offer. Taking into consideration that several of these papers may already be published or in the process of being published, we instructed the residents not to include any images already being used elsewhere. Fortunately (or unfortunately, depending on your perspective), we are all employees of the government, so there is no copyright to transfer our work.

Finally, the products here are presented by our residency training program, which adequately reflects the outstanding leadership provided over the past four years by COL Scott B. Shawen, MD. The past four years have seen unprecedented growth in the academic program and we are deeply indebted to his leadership. I am hopeful you will enjoy reading this collective sample of our program as much as I have. Thank you!

MAJ Melvin D. Helgeson, MD, MC, USA  
Orthopaedic Spine Surgeon  
Director, Orthopaedic Research  
Walter Reed National Military Medical Center  
Assistant Professor, Uniformed Services University of the Health Sciences  
Bethesda, MD

**Subpectoral Biceps Tenodesis: an Anatomical Study and Evaluation of At-Risk Structures**

<sup>1</sup>MAJ Jonathan F. Dickens, MD, MC, USA; <sup>1</sup>MAJ Kelly G. Kilcoyne, MD, MC, USA; <sup>1</sup>LCDR Scott M. Tintle, MD, MC, USN; <sup>1</sup>MAJ Jeffrey Giuliani, MD, MC, USA; <sup>1,2</sup>COL Richard A. Schaefer, MD, MC, USA; <sup>2</sup>CDR John-Paul Rue, MD, MC, USN

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**INTRODUCTION**

Pathology afflicting the proximal portion of the long head of the biceps has long been recognized as a common source of shoulder pain.<sup>11-3</sup> Numerous biceps tenodesis procedures have been described which secure the biceps tendon within<sup>1-5</sup> or inferior to the bicipital groove.<sup>6-8</sup> In 2005, Mazzocca, et al.<sup>7</sup> described an open subpectoral biceps tenodesis (OSPBT) technique securing the tendon below the intertubercular groove using an interference screw. The benefits of this technique include improved surgical efficiency, maintenance of the biceps length-tension relationship, removal of the tendon from the bicipital groove, and secure fixation.<sup>7,9,10</sup> Limited series have demonstrated favorable outcomes using the OSPBT technique,<sup>11</sup> with failure of fixation and persistent bicipital pain as the most commonly reported complications.<sup>12</sup>

The OSPBT with interference screw fixation has been increasingly performed for the treatment of multiple conditions of the proximal biceps tendon and associated shoulder pathology. There are numerous neurovascular structures in the proximal arm potentially at-risk during open subpectoral biceps tenodesis. Recently several studies have reported complications associated with the OSPBT technique.<sup>11-13</sup> Despite the potential for a catastrophic neurovascular injury during this procedure, there have been no studies that we are aware of in the English language which characterize the anatomic structures at risk in this relatively new surgical procedure. Additionally, the effect of arm rotation on the position of the neurovascular structures in the proximal arm has not been described. The purpose of this study was to define the anatomic relationships and at-risk structures during OSPBT and to quantify the effect of arm rotation on the position of the neurovascular structures of the proximal arm.

**METHODS**

After institutional review board approval, 17 whole intact fresh-frozen cadaveric upper extremities from nine donors (six male and three female) were studied. Nine right- and eight left-sided extremities were utilized, with eight paired specimens. The mean age at time of death was 77 years (range, 57 to 96 years). All specimens

thawed to room temperature before experimentation. No limbs had visual evidence of prior shoulder surgery or trauma. All shoulders maintained at least 160° of passive abduction and 160° of passive forward flexion.

**MAJ Jonathan F. Dickens MD, MC, USA**



MAJ Dickens attended Davidson College and received a bachelor of science in biology. Following his undergraduate education MAJ Dickens served as a platoon leader and operations officer in 161<sup>st</sup> Airborne Medical Support Battalion at Fort Bragg, NC. He then entered medical school at Indiana University School of Medicine where he was inducted into the *Alpha Omega Alpha* honor medical society. After medical school he began orthopaedic surgery residency at Walter Reed National Medical Center. MAJ Dickens completed the Walter Reed research fellowship under the mentorship of Dr. John-Paul Rue and Benjamin K. Potter. MAJ Dickens' research has focused on combat casualty care and sports medicine, with interests particularly in open calcaneus and elbow fractures, proximal biceps pathology, and shoulder instability. He is the principal or co-principal investigator for nine on-going projects and has received over \$350,000 in funding for his research efforts. His research has produced over 20 peer reviewed publications, five chapters, and more than 50 presentations to date. MAJ Dickens will complete the John A. Feagin Jr. Sports Medicine Fellowship at West Point in 2013 and he aspires to continue his research interests and sports medicine practice at an academic tertiary care hospital.

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The open subpectoral biceps approach, as described by Mazzocca, et al.<sup>7</sup> was performed by two investigators. As originally described, a pointed Hohman was placed directly against the lateral aspect of the humerus to retract the pectoralis major tendon proximally and laterally. A blunt Chandler was placed medially to retract the coracobrachialis and short head of the biceps. The retractors were positioned on the medial and lateral border of the humerus and centered at the point where the long head biceps intersected the inferior border of the pectoralis major. One cm proximal to the inferior border of the pectoralis major and centered at the inferior aspect of the bicipital groove, a guidewire was placed to identify and represent the proposed tenodesis site. The correct tenodesis position was confirmed in all specimens when the musculotendinous portion of the long head of the biceps rested at the inferior border of the pectoralis major tendon. A tenodesis screw was not placed in order to facilitate subsequent identification of neurovascular structures.

A careful dissection was then performed and structures were measured in a straight line from the at-risk structure to the pre-assigned reference points (described below) using standard digital calipers. Subcutaneous dissection was performed to identify the cephalic vein, medial brachial cutaneous nerve, medial antebrachial cutaneous nerve, and the intercostal brachial cutaneous nerve. Superficial structures were measured from the superior and inferior aspect of the 4-cm incision. Deep dissection was performed to identify the brachial artery, deep brachial artery, brachial vein, musculocutaneous nerve, median nerve, axillary nerve, and radial nerve. With the arm in the recommended position of 30° abduction and neutral rotation, the deep structures were measured from the tenodesis site and the nearest retractor. Additionally, the musculocutaneous nerve was measured with the arm in 45° of internal rotation and 45° of external rotation to determine the effect of rotation on the musculocutaneous nerve. Care was taken to minimize dissection and measure each structure in its native anatomic location.

## RESULTS

### Superficial Structures

With the arm at 30° of abduction and neutral rotation, the cephalic vein was 9.2 mm ± 6.1 mm lateral to the superior margin of the incision and 13.7 mm ± 5.8 mm lateral to the inferior margin of the incision. The medial brachial cutaneous nerve of the arm, medial brachial cutaneous nerve of the forearm, and intercostal brachial cutaneous nerves were 38.8 mm ± 4.2 mm, 33.8 mm ± 7.5 mm, and 45.9 mm ± 7.6 mm medial to the superior aspect of the incision, respectively.

### Deep Structures

With the arm at 30° of abduction and neutral rotation, the musculocutaneous nerve was 10.1 mm ± 3.3 mm medial to the tenodesis location and 2.9 mm ± 1.4 mm medial to the medially placed retractor (Table 1, Table 2). The radial nerve and deep brachial artery were 16.6 mm ± 5.9 mm and 14.5 mm ± 6.0 mm deep and medial to the tenodesis location, respectively. With the medial retractor in place, the radial nerve and deep brachial artery were 7.4 mm ± 3.0 mm and 5.7 mm ± 2.9 mm deep and medial to the retractor, respectively. The median nerve, brachial artery, and brachial vein were 27.6 mm ± 5.5 mm, 32.0 mm ± 5.7 mm, 32.5 mm ± 6.4 mm medial to the tenodesis site, respectively.

The distance between the musculocutaneous nerve and the tenodesis site was significantly less in internal rotation (8.1 mm ± 3.3mm) versus external rotation (19.4 mm ± 8.2 mm,  $p = 0.009$ ) (Table 3). In 18% of specimens (3/17), the musculocutaneous nerve was in direct contact with the medial retractor with the arm in an internally rotated position.

**Table 1. Distance from Tenodesis Site to Anatomic Structure in Neutral Rotation**

Structure	Mean (mm)	Standard deviation	Range (mm)
Musculocutaneous nerve	10.1	± 3.3	6-18
Axillary nerve	33.8	± 6.9	20-45
Median nerve	27.6	± 5.5	18-49
Radial nerve	16.6	± 5.9	10-32
Deep brachial artery	14.5	± 6.0	9-30
Brachial vein	32.5	± 6.4	19-45
Brachial artery	32.0	± 5.7	20-43

**Table 2. Distance from Retractor to Anatomic Structure**

Structure	Mean (mm)	Standard deviation	Range (mm)
Musculocutaneous nerve*	2.9	± 1.3	1-6
Median nerve*	20.9	± 6.1	13-31
Radial nerve*	7.4	± 3.0	2-12
Deep brachial artery*	5.7	± 2.9	1-10
Axillary nerve^	11.5	± 4.3	4-21

\* indicates medial retractor; ^ indicates lateral retractor

**Table 3. Variation in Musculocutaneous Nerve Relationship to the Tenodesis Site in Internal, Neutral, and External Rotation and the Number of Specimens in Contact with the Medial Retractor**

Position	Internal rotation	Range	Neutral rotation	Range	External rotation	Range	<i>p</i> value
Mean (mm)	8.1 ± 3.3	3.3 to 16.1	10.1 ± 3.3	6.4 to 14.5	19.4 ± 8.2	10.6 to 32.3	< 0.01
No. in direct contact	3		0		0		

Variations of musculocutaneous nerve innervations to the biceps and brachialis were observed and classified.<sup>14,15</sup> A type I biceps innervation, defined as a single motor branch from the musculocutaneous nerve was observed in 76% (13/17) of specimens. A type II biceps innervation, defined as two distinct motor branches from the musculocutaneous nerve was observed in 24% (4/17).<sup>14,15</sup> A type III innervation pattern, defined as three distinct musculocutaneous motor branches, was not observed.<sup>14,15</sup> The site of origin of the first branch was approximately half the distance between the lateral border of the acromion and the tip of the medial epicondyle of the humerus. A type I brachialis innervation was observed in 71% (12/17) and a type II brachialis was observed in 29% (5/17).<sup>14,15</sup> The first branch to the brachialis was consistently inferior to the branch to the biceps. There was no significant difference in measurements made by either investigator ( $p > 0.05$ ).

## DISCUSSION

Limited series have evaluated the outcomes following OSPBT.<sup>12,16,17</sup> The incidence of complications following open subpectoral tenodesis have been reported to be between 2% and 7%,<sup>12,18</sup> however, the incidence of musculocutaneous neuropraxia following OSPBT is not reported. Nho, et al.,<sup>12</sup> in the largest clinical series of OSPBT, reviewed the complications in 353 patients and found a 2.0% complication rate. Persistent bicipital pain and failure of fixation with a Popeye deformity were the most common complications, 0.57% each. One patient (0.28%) developed a musculocutaneous nerve neuropathy, presenting 10 days postoperatively with forearm numbness and weakness in elbow flexion and forearm supination six weeks postoperatively. Musculocutaneous nerve exploration six weeks following the index surgery demonstrated the nerve was in continuity and at six months postoperatively, the patient had complete resolution of the neurologic deficits.

Ma, et al.<sup>13</sup> reported another case of musculocutaneous nerve injury following OSPBT. Five months after OSPBT, the patient underwent musculocutaneous nerve exploration, demonstrating that the long head of the biceps tendon was transferred deep to the nerve, providing continued traction on the nerve. The nerve was found to be in continuity and following revision of the tenodesis with placement of the long head of the biceps superficial to the nerve, the patient regained full neurological function.

The proximity of the musculocutaneous nerve to the tenodesis site makes it the most at-risk structure during OSPBT. Additionally, the use of a deep, levering medial retractor places the nerve at increased risk of iatrogenic injury. The mean distance from the medial retractor to the musculocutaneous nerve with the arm in internal rotation was  $2.9 \text{ mm} \pm 1.4 \text{ mm}$ , and in 18% of specimens, the nerve was in direct contact with the retractor. Therefore, if the surgeon's preference is for medial retraction of the conjoint tendon and medial soft tissues, then a non-levering retractor with gentle retraction during the critical portions of the procedure is recommended to minimize iatrogenic injury to the musculocutaneous nerve.

External rotation of the arm significantly increased the distance from the tenodesis site by 11.3 mm. OSPBT is frequently performed following shoulder arthroscopy in either the lateral decubitus or beach chair positions. Mounted arm positioners may also be utilized to minimize the need for a surgical assistant and improve arthroscopic visualization. These techniques, however, limit the ability to position the extremity safely for OSPBT. We recommend positioning the arm in an externally rotated position to maximize the distance of the musculocutaneous nerve from the operative field.

Although no reported deep brachial artery or radial nerve injuries have been reported with OSPBT, the location of these structures as they course lateral and posterior into the spiral groove immediately posterior to the tenodesis is demonstrated and may represent a potential site of injury. Cox, et al.<sup>19</sup> reported that the radial nerve traversed the midpoint of the spiral groove  $14.7 \text{ cm} \pm 2.2 \text{ cm}$  distal to the greater tuberosity of the humerus at a point approximately 48% from the most proximal point of the humerus. Guse, et al.<sup>20</sup> found that the radial nerve passed posterior to the medial cortex of the humerus at a point 41% of the humeral length ( $12.4 \text{ cm} \pm 1.2 \text{ cm}$  from the acromion) and exited the spiral groove lateral to the lateral cortex of the humerus at a point 58% of the humeral length ( $17.6 \text{ cm} \pm 1.7 \text{ cm}$  from the tip of the acromion). A medially placed retractor that penetrates the fascia of the short head of the biceps may come in direct contact with the radial nerve and deep brachial artery as they course into the spiral groove. Additionally, inadvertent bicortical drill penetration during tenodesis preparation may compromise the structures within the spiral groove. In all specimens, the structures of the spiral groove were within 1.5 cm and at the level of the proposed tenodesis site. Drilling techniques to avoid bicortical penetration of the proximal humerus should be employed. Knowledge of the proximity and location of these structures may prevent iatrogenic injury.

There have been no cases of axillary nerve or artery damage following OSPBT. Although unlikely, a retractor placed on the lateral humerus under the pectoral major muscle may jeopardize the axillary nerve if vigorously retracted. In this series, the axillary nerve was a mean  $11.5 \text{ mm} \pm 4.3 \text{ mm}$  from the retractor. Care should be taken to limit deep placement and excessive force during lateral retraction.

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## Complications and Reoperations Associated with Combat-Related Tibial Plafond Fractures

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### INTRODUCTION

Since the onset of the war on terror, nearly 46,000 U.S. military personnel have been listed as wounded in action.<sup>1</sup> Recent analyses have shown that a majority of the sustained injuries are the result of explosions from mortars, rockets, and improvised explosive devices (IEDs).<sup>2</sup> Because of effective modern body armor and Kevlar that protect the torso and head relatively well, it is the extremities that absorb most of the energy from blast mechanisms. Consequently, a high number of these extremity injuries are long bone fractures.<sup>3,4</sup>

Combat-related injury to the tibia has been the subject of many articles in the literature over the past several years. However, while there is a significant amount of literature regarding tibial shaft fractures, there is a relative paucity of literature on periarticular injuries to the tibia among wounded soldiers. Open fractures secondary to blast trauma have a high incidence of complications and open periarticular injuries portend an even worse prognosis.<sup>5-8</sup> Fractures to the tibial plafond among the war-wounded have not been a subject directly addressed since the recent war efforts began. The complications and outcomes of these devastating injuries have yet to be elucidated in the literature. In this study, we aim to report the experience at our institution with these injuries, as well as identify and describe the complications and re-operations following treatment of combat-related tibial plafond fractures.

### METHODS

After obtaining approval from our institutional review board (IRB), we performed a retrospective review of all patients with tibial plafond fractures treated at one of our three institutions, Walter Reed Army Medical Center, National Naval Medical Center–Bethesda, or Walter Reed National Military Medical Center, between 2003 and 2012. Only patients who were active duty military at the time of injury, sustained the pilon fracture as a result of combat operations, and had definitive care rendered at one of our three affiliated institutions met criteria for inclusion in our study. Patients who sustained fractures secondary to non-combat etiology or who were non-active duty military were excluded.

Based on our inclusion criteria, a total of 63 fractures in 62 patients were identified (one patient sustained bilateral pilon fractures). Patient demographic data to include age and gender were obtained. For each fracture, we recorded mechanism of injury (blast, gunshot, motor

vehicle collision, fall, etc.), open versus closed fracture, associated injuries, time from injury to definitive fixation, type of definitive fixation (open reduction with internal fixation (ORIF), external fixation, or hybrid fixation), number of unplanned re-operations after definitive fixation, reason for any re-operations, and total length of follow-up. Additionally, open fractures were classified based on the Gustilo-Anderson classification<sup>9,10</sup> and recorded according to the type of definitive soft tissue coverage used in relevant cases.

Patients were then stratified into three cohorts based on type of fixation: ORIF, external fixation, or hybrid fixation. These cohorts were subsequently compared with the previously described variables to determine any difference in complication rate based on type of fixation.

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LCDR Grijalva grew up in Tucson, AZ. Shortly after graduation, he enlisted in the Navy becoming a hospital corpsman. He spent seven years enlisted prior to receiving his bachelor of science degree from Excelsior College in Albany, NY. He received his doctor of medicine from the Uniformed Services University of the Health Sciences. Shortly after completing his orthopaedic internship at the National Naval Medical Center, he transferred to Camp Lejeune, NC and was assigned to 2<sup>nd</sup> Marine Division. During his stay, he was attached to 2<sup>nd</sup> Amphibian Assault Battalion, and 1<sup>st</sup> Battalion, 8<sup>th</sup> Marines. He completed a tour overseas in support of Operation Iraqi Freedom. Currently, he is an orthopaedic resident at the Walter Reed National Military Medical Center assigned to The Naval Medical Research Center for his research fellowship. His research has focused on heterotopic ossification (HO) and the effects of HO prophylactic agents on wound healing. He plans to apply for an orthopaedic trauma fellowship following residency graduation.

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## RESULTS

### All Fractures (Tables 1 & 2)

The average age of subjects was 26.8 years. Fifty-five of the 63 fractures were secondary to blast mechanism of injury; four secondary to falls; three secondary to gunshot wounds; and one secondary to motor vehicle collision. Twenty-nine of the fractures were open while

34 were closed. The average time to fixation from injury among all fractures was 19 days. Forty-three of the 63 ankles exhibited residual pain at long-term follow-up (greater than 12 months).

The average number of unanticipated re-operations was 1.82 among all patients. Infection was the reason for revisit to the operating room in 24% (15/63) of fractures; wound dehiscence in 20.6% (13/63); symptomatic hardware requiring removal in 9.5% (6/63); and 23.8% of fractures returned to the operating room for elective amputa-

**Table 1. All Fractures**

	No. of pilons N = 63	ORIF N = 41 (65.1%)	Ex-fix N = 11 (17.5%)	Hybrid N = 11 (17.5%)	p value
Age in years (mean $\pm$ standard)	26.8	27.8 $\pm$ 1.03	26.2 $\pm$ 2.00	23.6 $\pm$ 2.00	0.1
MOI					0.6
Fall	4	4 (100%)	0 (0%)	0 (0%)	
GSW	3	2 (66.67%)	1 (33.33%)	0 (0%)	
Blast	55	34 (61.8%)	10 (18.2%)	11 (20%)	
MVC	1	1 (100%)	0 (0%)	0 (0%)	
Open/closed					0.03*
Open	29	14 (48.3%)	8 (27.6%)	7 (24.1%)	
Closed	34	27 (79.4%)	3 (8.8%)	4 (11.8%)	
Time of definitive fixation (days)	19.06	15.02 $\pm$ 8.9	30.55 $\pm$ 17.5	22.64 $\pm$ 25	0.0068*
Total no. of re-operations	1.82	2.07 $\pm$ 3.62	1.81 $\pm$ 2.75	0.91 $\pm$ 0.70	0.56
Pain at > 12 months	43 (68.3%)	27 (62.8%)	7 (16.3%)	9 (20.9%)	0.56

ORIF, open reduction and internal fixation; ex-fix, external fixation; MOI, mode of injury; GSW, gun shot wound; MVC, motor vehicle collision;

\*, not significant

**Table 2. All Fractures Complications**

	No. of pilons N = 63	ORIF N = 41 (65.1%)	Ex-fix N = 11 (17.5%)	Hybrid N = 11 (17.5%)	p value
Infection	15 (23.8%)	10 (66.7%)	3 (20.0%)	2 (13.3%)	0.87
Wound complications	13 (20.6%)	9 (69.2%)	3 (23.1%)	1 (7.7%)	0.5
Symptomatic hardware	6 (9.5%)	6 (100%)	0 (0%)	0 (0%)	0.1
Amputation	15 (23.8%)	9 (60%)	4 (26.7%)	2 (13.3%)	0.5
Union	57 (90.5%)	38 (92.7%)	10 (90.9%)	9 (81.2%)	0.5
Average follow-up (months)	23.4	23.1 $\pm$ 17.7	30.3 $\pm$ 25.1	17.7 $\pm$ 7.4	0.2

ORIF, open reduction and internal fixation; ex-fix, external fixation

tion secondary to persistent residual pain and limited function. Overall, 90.5% (57/63) of fractures exhibited evidence of radiographic union regardless of complications. We defined union as osseous bridging at six months post fixation.

### Open Fractures (Table 3)

Re-operations were further stratified into open fractures alone and analyzed. All 20 open fractures were classified at Gustilo-Anderson type III fractures. Seventeen were type IIIA, 10 were type IIIB, and two were type IIIC. A total of 20.1% (6/29) required a flap (free or rotational) for adequate soft tissue coverage. The average number of unanticipated re-operations for open fractures was 2.17. Thirty-one percent (9/29) were secondary to infection; 31% (9/29) for wound complications; 13.8% (4/29) for symptomatic hardware; and 27.6% (8/29) underwent elective amputation. Eighty-two percent (24/29) of the open fractures showed evidence of radiographic union regardless of complications.

### Complications Based on Different Treatment Modalities (Tables 1 to 3)

Of the total 63 fractures, 65% (41/63) were treated with ORIF alone, 17.5% (11/63) with external fixation alone, and 17.5% (11/63) with hybrid fixation. There was no significant difference with respect to age between the three treatment modalities ( $p = 0.1$ ) nor was there a significant difference among mechanisms of injury ( $p = 0.6$ ). With respect to open versus closed fractures and treatment, 48.3% (14/29)

of open pilons were treated with ORIF alone, while the other 15 underwent external fixation alone or hybrid fixation, 27.6% (8/29) and 24.1% (7/29), respectively. Of the 34 closed fractures, 79.4% (27/34) underwent ORIF while 8.8% (3/34) and 11.8% (4/34) underwent external fixation and hybrid fixation, respectively. This difference in treatment modality based on open versus closed fractures was significant ( $p = 0.03$ ).

The difference in time to definitive fixation was also shown to be significant between treatment cohorts, with ORIF alone being performed an average of 15 days (+/- 8.9 days) post injury, external fixation alone at 30.55 days (+/- 17.5 days), and hybrid fixation at 22.6 days (+/- 25 days) ( $p < 0.01$ ). Differences in total number of re-operations and ankles with pain at long-term follow-up were not found to be significant between treatment cohorts. There was no significant difference in reasons for re-operations based on treatment type. Among open fractures stratified based on treatment, the only significant difference ( $p = 0.042$ ) was found with respect to utilization of a flap for soft tissue coverage, as 33.3% (2) of the total six flaps were performed in the ORIF group and 66.6% (4) were in the external fixation groups.

### DISCUSSION

Because of recent improvements in rehabilitative care of combat casualties, limb salvage for high-energy traumatic injury to the lower limb has become more practical.<sup>11-13</sup> However, the functional outcome and complications following limb salvage for combat-related tibial

Table 3. Open Fractures

	No. of open pilons N = 29	ORIF N = 14 (48.3%)	Ex-fix N = 8 (27.6%)	Hybrid N = 7 (24.1%)	p value
<b>Gustilo-Anderson classification</b>					0.08
<b>IIIA</b>					
<b>IIIB</b>	17 (59%)	10 (58.8%)	2 (11.8%)	5 (29.4%)	
<b>IIIC</b>	10 (34%)	4 (40%)	4 (40%)	2 (20%)	
	2 (6.9%)	0 (0%)	2 (100%)	0 (0%)	
<b>Infection</b>	9 (31%)	5 (55.6%)	2 (22.2%)	2 (22.2%)	0.86
<b>Flap for soft tissue coverage</b>	6 (20.1%)	2 (33.3%)	4 (66.7%)	0 (0%)	0.042*
<b>Total no. of re-operations</b>	2.17	2.85 ± 2.54	2.25 ± 3.10	0.71 ± 0.75	0.18
<b>Wound complications</b>	9 (31%)	6 (66.7%)	2 (22.2%)	1 (11.1%)	0.37
<b>Symptomatic hardware</b>	4 (13.8%)	4 (100%)	0 (0%)	0 (0%)	0.08
<b>Amputation</b>	8 (27.6%)	4 (50%)	3 (37.5%)	1 (12.5%)	0.60
<b>Union</b>	24 (82.8%)	12 (50%)	7 (29.2%)	5 (20.8%)	0.66

ORIF, open reduction and internal fixation; ex-fix, external fixation; \*, not significant



plafond fractures has yet to be elucidated. Our goal in this study was to identify risk factors for return trips to the operating room and describe our institution's experience with these difficult injuries. Additionally, we aim to compare different treatment options with respect to these complications, potentially in order to make improved surgical recommendations to patients with complex pilon fractures.

A number of previous studies have profiled the complications associated with pilon fractures in the civilian population. The primary reasons for re-operations in these studies are deep infection or elective amputation for persistent pain and/or disability.<sup>14-17</sup> Sirkin, et al. evaluated outcomes and complications of surgical management of pilon fractures in 56 patients, finding 5% and 11% infection rates in closed and open fractures, respectively.<sup>15</sup> A later study by Conroy, et al. of 32 open pilons showed an 18% infection rate and a 6.2% amputation rate.<sup>16</sup> Boraiah, et al. found an infection rate of 3% and an amputation rate of 2% among 59 pilon fractures.<sup>17</sup> Our study, in comparison, revealed a much higher complication rate. Among all pilon fractures in our patient population, we found an overall infection rate of 24% and an amputation rate of 20.8%. In open fractures alone, the infection and amputation rates were 31% and 27%, respectively. Additionally, 20% of the open fractures were Gustilo-Anderson type IIIB fractures requiring rotational or free flap soft tissue coverage. This complication rate is significantly higher than that reported in the civilian literature.

The reason for this stark difference in complication rates is not completely clear. We hypothesize, as previous studies on the war-wounded have also indicated, that the high-energy nature of injuries sustained in combat are the foundations for subsequent complications in this patient population.<sup>7,18,19</sup> The majority (55 of 63) of the fractures in our study were sustained secondary to blast mechanisms; this is in stark contrast to the mechanisms of injury for tibial plafond fractures in the civilian population, which are usually secondary to falls or motor vehicle collisions.

We feel that the high complication rate in our study reflects not the treatment actually rendered, but rather the severity of the initial injury. Despite this high complication rate, over 90% of the pilon fractures in our patient population showed evidence of radiographic union at six months follow-up, indicating that it is soft tissue insult that likely causes the increased re-operation rate.

Since Ruedi and Allgower's landmark study in 1973,<sup>20</sup> surgical management of pilon fractures has been the mainstay of treatment for these injuries. They showed promising outcomes with operative intervention, as opposed to the previous dogma of non-operative management.<sup>21</sup> Numerous reports have since documented various treatment options for pilon fractures, to include ORIF, definitive external fixation, and hybrid fixation.<sup>15,22-28</sup> Bone, et al. showed generally favorable outcomes in patients treated with external fixation in combination with ORIF using small plates and percutaneous screws near the articular surface to limit soft tissue dissection.<sup>24</sup> In a prospective study of 26 patients comparing ORIF alone with hybrid treatment, Tornetta, et al. revealed overall similar outcomes and less soft tissue complications in the hybrid cohort.<sup>28</sup> A later study by Anglen, et al. showed results inconsistent with earlier reports on the improved outcomes and complication rates of hybrid fixation. They found lower clinical scores, slower return to function, a higher rate of complications, more infections, and a higher incidence of malunions and nonunions in 63 patients treated with hybrid fixation.<sup>22</sup> As a result of these disparities in the literature, there remains no clear consensus on the ideal form of management of these injuries.

Our study showed no significant difference in re-operation rates between treatment cohorts. The only significant differences with respect to the different treatment protocols were time to definitive fixation and whether the fracture was an open or closed injury. Patients treated with an external fixator, either alone or using a hybrid technique, were much more likely to have a lengthier delay in definitive treatment from initial injury. This is likely due to the fact that more severe injuries involved greater soft tissue insult, resulting in a lack of feasibility of adequate internal fixation; these fractures consequently were more likely to benefit from some form of definitive external fixation. We believe that for similar reasons, open fractures were more likely to be treated definitively with external fixation, as open injuries portended to more significant soft tissue compromise.

Although not found to be statistically significant, we observed a much higher proportion of patients with deep infection had been treated with ORIF alone (66.7%), in contrast to external fixation alone (20%) and hybrid fixation (13.3%). If our sample size had been larger, we may have found a statistical difference, but it appears necessary to highlight this difference nevertheless.

We recognize that our study has several limitations. First, the retrospective nature of the study limits the extent to which we can make conclusions from our results. Second, our study lacks a detailed profile of outcomes of fractures, as we did not perform objective measures or surveys to assess functionality and long-term follow-up. Finally, our small sample size limits the power of our study and the potential to make sound conclusions. Our study, however, is the first to describe the complexity in treatment of tibial plafond fractures among the combat-wounded population. Further studies are needed to make definitive recommendations with regard to these injuries.

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## Reported Concussion Rates for Three Division I Football Programs: an Evaluation of New NCAA Concussion Policy

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### INTRODUCTION

Concussions are common injuries in young athletes, but the diagnosis and treatment remains a challenge.<sup>1,2</sup> On April 29, 2010 the NCAA released a new concussion management plan which provided a best practices model for concussion treatment. The purpose of this study is to determine the number of concussions that occurred on three collegiate Division I football teams during the two most recent seasons, 2009-2010 and 2010-2011, in order to compare the incidence of concussion before and after the implementation of these new policies. We hypothesized that the number of reported concussions and concussion rate would be consistent among the three teams, and from season to season.

### MATERIALS AND METHODS

Injury surveillance reports were reviewed for three NCAA Division I football teams: the United States Military Academy, the United States Naval Academy, and the United States Air Force Academy. All subjects included in the study were males between the ages of 18 to 22, who were on the varsity football roster at their respective institution for the 2009-2010 season and/or the 2010-2011 season.

Regardless of institution, and in accordance with the NCAA Injury Surveillance System definition of a reportable injury, a concussion was defined as an injury that occurred as a result of participation in organized intercollegiate practice or competition that required medical attention by a team certified athletic trainer or physician that resulted in restriction of student athlete play for one or more calendar day after injury, and was documented in the injury report by the athletic trainer as a concussion.<sup>3</sup>

The primary outcome of interest in the current study was the annual incidence rate of concussion injuries per 1,000 athlete-exposures during the study period. Incidence rates are calculated by dividing the total number of injuries observed in a population by a measure of exposure or person-time at risk to injury.<sup>4</sup> The annual incidence rates of concussion for the 2009-2010 and 2010-2011 seasons were calculated, along with the 95% confidence intervals, by dividing the total number of concussions by the total number of athlete-expo-

sure and multiplying by 1,000. The incidence rates for the 2009-2010 and 2010-2011 seasons were compared by calculating the incidence rate ratio and 95% confidence interval using the 2009-2010 season as the referent category. Similarly, we calculated the annual incidence rate with 95% confidence intervals for each team for the season prior to, and the season following implementation of the new NCAA concussion management policy. The incidence rates for the 2009-2010 and 2010-2011 seasons were also compared within each team by calculating the incidence rate ratio and 95% confidence interval using the 2009-2010 season as the referent category. All statistical analyses were completed using STATA/SE software version 10.1 (StataCorp, College Station, TX).

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MAJ Kilcoyne obtained his doctor of medicine degree at the University of Maryland on a health profession scholarship. He was selected for orthopaedic residency at Walter Reed Army Medical Center and is currently in his final year. During his residency MAJ Kilcoyne co-authored over 15 peer-reviewed articles and three book chapters and completed over 45 research presentations at national and local research meetings. His research interest has primarily focused on sports injuries, however, he has also co-authored articles or chapters on war-related injuries. In addition to several research grants, MAJ Kilcoyne's research projects have also won awards; he was a finalist for the Founders Award at the SOMOS annual meeting. He has been selected to complete a shoulder and elbow fellowship at The Johns Hopkins University upon graduation.

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## RESULTS

The incidence rate and number of reported concussions increased at each institution from the 2009-2010 season to the 2010-2011 season after the implementation of the new NCAA concussion management policy. Overall, the incidence rate across all three institutions studied was two times higher in 2010-2011, when compared to 2009-2010 (IRR 2.04, 95% CI: 1.20, 3.55,  $p = 0.005$ ). Similar increases were observed across all three institutions when they were examined independently.

## DISCUSSION

The combined incidence rate of reported concussions from three Division I football programs for the 2010-2011 season doubled from the previous season after the implementation of new NCAA policies on concussion management. The two-fold increase in the combined concussion incidence rate between the two consecutive seasons is striking. While the institution of a more formalized concussion plan on the part of medical staff is one possible factor, the largest difference may have been from previous under-recognition and under-reporting on the part of players and coaches prior to the new policy. Additional studies are needed to determine the true incidence of concussions in Division I football as awareness continues to increase among healthcare professionals, coaches, and athletes. Furthermore, as similar policies are implemented across the United States, systematic evaluations are needed to assess fully whether these policy changes are reducing the impact of concussion among athletes.

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# BMP Regulation of Mesenchymal Progenitor Cells in Regenerating Muscle Tissue

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## INTRODUCTION

Post-traumatic heterotopic ossification (HO) is condition characterized by ectopic bone formation during muscle regeneration that occurs following orthopaedic injury. However, traumatic HO occurs much more frequently in patients who have experienced high-energy extremity injury and polytrauma. More than 60% of patients who sustained extremity blast-related trauma during Operation Enduring Freedom and Operation Iraqi Freedom were diagnosed with HO within one year of their injuries.<sup>1</sup> Ectopic bone formation can lead to impaired prosthetic fitting, skin breakdown, decreased joint range of motion, and painful joint range of motion, leading to significant morbidity to the men and women of our armed forces. Given the high incidence of this disease in the war-wounded population, there is a substantial need to characterize the microenvironment of the injured muscle tissue to identify factors that may be dysregulating the process of regeneration and to propose new treatment options that will minimize the formation of ectopic bone. The roles of bone morphogenetic proteins (BMPs) are of particular interest, as they have recently been associated with a variety of functions in healthy and injured muscle tissue,<sup>2,3</sup> and are also capable of inducing osteogenic differentiation.

Our laboratory has previously identified a population of multipotent mesenchymal progenitor cells (MPCs) that are present in regenerating muscle tissue after high-energy trauma.<sup>4</sup> Although these cells are capable of osteogenic differentiation, they also perform many of the functions that are characteristic of bone marrow-derived mesenchymal stem cells, specifically, promoting tissue regeneration by enhancing angiogenesis, regulating fibrogenesis, and modulating inflammation.<sup>5</sup> The effect of BMPs secreted by the traumatized muscle tissue on MPC function and the precise role of these cells in ectopic bone formation is unknown. Therefore, the goal of this study is to better define the interactions between MPCs and the BMPs that are present in the traumatized muscle tissue environment. Our overall hypothesis was that the BMPs dysregulate the pro-regeneration functions of the MPCs to promote pathological wound healing over muscle regeneration. We propose the following specific aims:

- 1) To compare the gene- and protein-level expression of BMPs that were expressed in the traumatized and uninjured muscle tissue.
- 2) To determine the effect of the secreted BMPs on the anti-fibrotic properties of the traumatized muscle-derived MPCs.

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## METHODS

Muscle samples were obtained from the zone of injury following blast trauma according to an IRB-approved protocol. Control muscle tissue was obtained from patients undergoing elective orthopaedic anterior cruciate ligament reconstruction with hamstrings autograft. After the tendons were harvested with a tendon stripper, the remaining muscle on the tendon was removed. Muscle fragments approximately 0.2 cc in volume were taken from all control and traumatized muscle samples. One fragment was homogenized to determine the BMP gene expression with qPCR. Protein-level measurements of BMP were determined using the other fragments by determining their wet-weight and placing them in 12-well transwell plates containing 2 ml of growth medium (DMEM supplemented with 10% fetal bovine serum and 1% penicillin and

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streptomycin). The growth medium was collected every three days, and the concentration of BMP-2, -4 and -6 was measured in the collected medium with ELISAs (R&D Systems).

MPCs were harvested from the debrided muscle samples by following a previously described protocol.<sup>4</sup> The protocol for extracting muscle-derived multiprogenitor cells was based on a modification of previous work in isolating mesenchymal stem cells that was performed in our laboratory. Fat, fascia, other connective tissue, and necrotic tissue were dissected away from the healthy margin of the debrided muscle sample. Approximately 0.2 cc of the remaining muscle tissue was processed for cell extraction. The tissue was washed three times in Hanks' Balanced Salt Solution (Gibco, Carlsbad, California) and then was extensively minced in a 10-cm culture dish containing Dulbecco's Modified Eagle Medium (Gibco) and 3· penicillin/streptomycin/Fungizone (Gibco) until it could pass through the tip of a 25-mL serological pipette (Falcon; BD Biosciences, San Jose, California). The minced tissue was transferred to a 50-mL conical vial with digestion medium containing Dulbecco's Modified Eagle Medium, 3· penicillin/streptomycin/Fungizone, and 0.5 mg/mL collagenase type 2 (Worthington Biochemical, Lakewood, New Jersey). The tissue slurry was agitated gently at 37° C for two hours, and the resulting digest was filtered through a 40-mm cell strainer (Falcon), pelleted by means of centrifugation, resuspended in growth medium (Dulbecco's Modified Eagle Medium with 10% fetal bovine serum; Gibco) and 5· penicillin/streptomycin/Fungizone, and then plated onto tissue culture polystyrene (150-cm<sup>2</sup> flask; Falcon). The cells were incubated at 37° C in a 5% CO<sub>2</sub>-humidified cell incubator for two hours and then were extensively washed with Hanks' Balanced Salt Solution before fresh growth medium was added with 3· penicillin/streptomycin/Fungizone. Once multiprogenitor cell colony forming units were observed, the concentration of penicillin/streptomycin/Fungizone was lowered to 1·. Cell confluence was obtained after approximately two weeks. The cell cultures were routinely passaged at 80% to 90% confluence and split 1:4.

The cells were seeded in 12-well plates in growth medium (Dulbecco's Modified Eagle Medium with 10% fetal bovine serum; Gibco) and 5· penicillin/streptomycin/Fungizone, allowed to adhere overnight, and then treated with either BMP-4 over the range of concentrations that was observed during the BMP release kinetics experiment. The cell supernatants were collected after 72 hours and one week. These time points were chosen because they correlated with the largest spike in concentration of BMP-4 in the original explant data. The supernatant was stored in a -80° C freezer. The effect of the BMPs on the ability of MPCs to regulate fibrogenesis was assayed by measuring the concentration of matrix metalloproteinase-1 (MMP-1) in the cell supernatants with ELISA (DuoSet ELISA Development System, R&D Systems).

MPCs isolated from traumatized muscle tissue were exposed to BMP-4 in concentrations of 5 ng/ml, 50 ng/ml, and 200 ng/ml in growth medium. Quantitative real-time polymerase chain reaction (qRT-PCR) was performed on cells exposed to BMP-4 using an RT<sup>2</sup> profiler PCR array for the human osteogenesis pathway. The MPCs were exposed to BMP-4 for a total of seven days and then lysed with Trizol. RNA was extracted using the RNeasy Minikit, and RNA quality was assessed using the ND 1000 Nanodrop Spectrophotometer prior to performing the qRT-PCR assay.

## RESULTS

We measured substantial gene-level expression of BMP-2, BMP-4, BMP-5 and BMP-6 in control muscle. The gene expression of BMP-2 and BMP-4 was significantly lower in the traumatized muscle tissue ( $p < 0.015$ , Student's *t*-tests), and this was contrary to what we initially expected. Protein-level expression of BMP peaked at one week in culture, but was detected in the explant supernatants for up to 21 days, as determined by ELISA, indicating that BMP-4 is expressed by muscle and its surrounding microenvironment as it degenerates, despite decreased translation. We also measured a significant increase in the amount of MMP-1 expressed by the MPCs that were cultured with BMP-4 at concentrations of 50 ng/ml and 200 ng/ml ( $p < 0.05$ , Repeated Measures ANOVA with  $n = 3$ ). Cartilage oligomeric matrix protein (COMP) was significantly up-regulated in cells exposed to 200 ng of BMP 4 for seven days when compared with controls ( $p < 0.05$ ). Conversely, tumor necrosis factor (TNF) was significantly down-regulated in cells exposed to 200 ng of BMP 4 for seven days when compared to controls ( $p < 0.05$ ), as measured by PCR.

## DISCUSSION

It is unclear whether traumatized muscle-derived MPCs have a direct or an indirect role in the formation of ectopic bone following high-energy extremity injury. The MPCs are capable of osteogenic induction; although other wound healing cells have also been implicated in the formation of HO. MPCs which have a potent response to BMPs in vitro do not have the same robust response in vivo. However, cells expressing vascular endothelial markers do contribute to the generation of a fibroproliferative, chondrogenic, and ossification in the fibrodysplasia ossificans progressiva (FOP) model.<sup>6</sup> Alternatively, the MPCs may also play an indirect role in promoting HO formation by changing the biochemical microenvironment to favor fibrosis and ectopic osteogenesis over muscle regeneration.

BMP-4 gene expression is down-regulated in response to injury, which was contrary to our original hypothesis. There is a significant amount of data from the literature regarding FOP, a congenital disorder leading to ectopic bone formation, indicating that BMP-4 is dysregulated.<sup>11,12</sup> Specifically, the negative feedback loop normally present in BMP-4 signaling is dysregulated, leading to a hyperresponsiveness to ligand exposure.<sup>13</sup> We therefore continued to investigate protein expression with ELISA data for BMP 4, and there was an increase in BMP-4 accumulation in the explant. BMP-4 could be released from the extracellular matrix during the early remodeling stages of wound healing. BMP-4 has previously been shown to promote anti-fibrotic muscle regeneration by down-regulation of TGF- $\beta$ 1 and up-regulation of MMPs.<sup>7</sup> We found that increased BMP concentration resulted in a corresponding increase in MMP-1 concentration, which is an extracellular matrix protein essential in matrix remodeling in wound healing and decreased fibrosis.<sup>8</sup> These results indicate that MPCs may contribute to local wound healing, even when exposed to BMP-4. However, chronic exposure to BMP-4 may have deleterious effects.

Additionally, when MPCs were exposed to BMP-4, there were significant changes in the osteogenic gene expression profile. BMP-4 exposure led to an up-regulation of COMP expression. COMP is an extracellular matrix calcium binding protein found in cartilage and is essential for normal endochondral ossification and longitudinal development of the physis. COMP mutations are associated with several

different growth abnormalities resulting in rhizomelic shortening, including multiple epiphyseal dysplasia and pseudoachondroplasia.<sup>9</sup> There was also a significant decrease in TNF. BMP-4 has previously been shown to have an anti-apoptotic effect on mesenchymal pluripotent cells via a suppression of TNF-mediated apoptosis,<sup>10</sup> indicating that the BMP-4 found in traumatized muscle tissue may exert a protective effect on the MPCs resident in the tissue.

Our results suggest that there may be a window following injury, during which functional muscle regeneration is promoted via BMP-4 release from the muscle tissue. In cases of severe injury, the initial stress response is exaggerated, leading to a longer period of time when MPCs are present in the wound bed. In this setting of prolonged or amplified muscle regeneration, MPCs may also survive longer via the anti-apoptotic effect of BMP-4; the regenerative response may shift to form fibrotic tissue. This, in turn, generates an environment with the up-regulation of COMP that is susceptible to HO formation.

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## Application of the Orthoplastic Reconstructive Ladder to Preserve Lower Extremity Amputation Length

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### INTRODUCTION

Since the Global War on Terror commenced in 2001, over 1,200 U.S. service members have sustained in excess of 1,600 amputations.<sup>1</sup> A primary goal in lower extremity amputation management is limb length preservation and the maintenance of viable joints.<sup>2</sup> Energy expenditure during ambulation directly correlates with residual limb length, preservation of functional segments, and preservation of a stable joint.<sup>3,4</sup> These tenants directly contribute to the resumption of a more normal gait pattern, increased functional capacity, and decreased energy expenditure.<sup>5</sup> The ability to preserve limb length, functional segments, and joints is significantly dependent on achieving soft tissue coverage. The reconstructive ladder serves as a guide for surgeons to employ the best possible tactic based on the soft tissue defect.<sup>6</sup>

The aim of this paper is to illustrate through case reports the application of the reconstructive ladder for preserving residual limb length in lower extremity traumatic amputations. Modalities employed to achieve soft tissue coverage for preservation of limb length are presented. Concepts from the reconstructive ladder<sup>7</sup> were utilized to determine the optimal method for achieving soft tissue coverage and thus preserving amputation residual length.

### CASE 1

A U.S. soldier injured by an improvised explosive device (IED) blast sustained multisystem trauma, including a traumatic transtibial amputation (Figure 1). The level of bony amputation was distal to the tibia tubercle and the knee extensor mechanism was intact. However, the zone of soft tissue injury extended well above the knee joint. There was inadequate soft tissue to achieve primary closure, to perform an effective myodesis or myoplasty, or to provide a well-padded residual limb. Treatment options included revising the amputation to a knee disarticulation or a transfemoral amputation. To preserve length, limb segment, and a functional joint, the patient underwent soft tissue coverage with a latissimus free flap, followed by split thickness skin coverage. The patient progressed to ambulation with a functional below-knee amputation without assist devices.

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### CASE 2

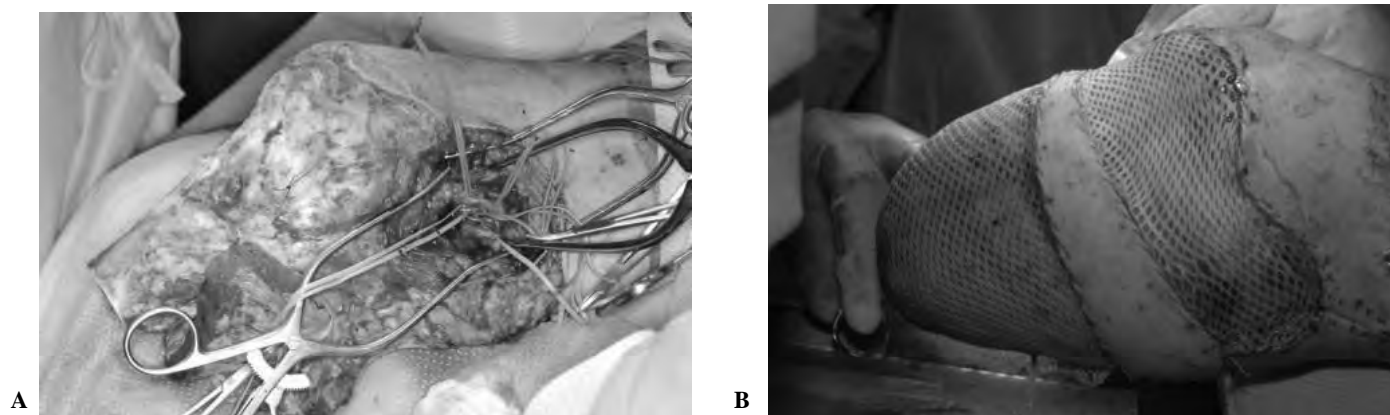
ACell (ACell, Inc, Columbia, MD) is a porcine-derived biomatrix scaffold that assists in tissue regeneration. In this case, ACell was employed to regenerate granulation tissue over avascular retinaculum and patellar ligament. Once adequate granulation tissue developed, it was amenable for split thickness skin grafting (STSG). This patient, who additionally sustained significant soft tissue injuries to his left lower extremity and bilateral upper extremities, is currently able to ambulate on his below-knee prosthesis without assist devices.

**LCDR Joseph Anthony O'Daniel, MD**



LCDR O'Daniel graduated with merit from the U.S. Naval Academy in 1996. Upon graduation, he served for five years as a surface warfare officer, stationed in Sasebo, Japan and Virginia Beach, VA. In 2001, he took a health professions scholarship and returned home to attend the University of Louisville School of Medicine. After graduating in 2005, he did his internship at Bethesda National Naval Medical Center and then served as the Battalion Surgeon for Naval Mobile Construction Battalion (NMCB) One in Gulfport, MS for two years. While there, NMCB One deployed to Okinawa, Japan for six months and then to Camp Ramadi, Iraq for seven months. Lcdr O'Daniel returned for his orthopaedic residency at Walter Reed Bethesda from 2008 through June 2013. Upon graduation, he and his wife Andrea have received orders to Great Lakes Naval Station just north of Chicago, IL.





**Figure 1.** (A) Use of free flaps and skin grafting. This wounded soldier sustained a traumatic below-knee amputation with significant soft tissue loss about his right knee and residual proximal tibia from a blast injury. (B) A latissimus free flap allowed coverage over the distal tibia, with split thickness skin grafting, as seen here.

## DISCUSSION

The preservation of residual limb length in lower extremity amputations is critical to optimize prosthetic fitting and to allow maximal function for ambulation. Although intuitive, a longer limb, preservation of joints, and more distal segments allow for decreased energy requirements when ambulating. As described by Waters, et al., amputees use a significantly higher energy requirement with a transfemoral versus a transtibial amputation.<sup>3</sup>

With respect to mangled extremity trauma, the soft tissue envelope is the greatest deciding factor for limb salvage versus amputation<sup>8</sup> and soft tissue availability often determines amputation salvage level. A goal in amputation surgery is the provision of a well padded durable surface for prosthetic fitting. Careful handling and preservation of all available soft tissue is critical. However, it is also important to keep in mind that muscle balancing is critical in order to preserve a functional residual limb. Often, soft tissue coverage can only be achieved by significant shortening of the residual bone. In this regard, salvage of limb length can be accomplished utilizing the same reconstruction principles for salvage of an intact limb.

Multiple authors, including Levin, have described the reconstructive ladder principal whereby soft tissue reconstruction begins at the lowest possible rung and advances upward based on the need and degree of soft tissue injury.<sup>9-11</sup> This traditional ladder approach includes techniques such as local wound care, primary closure, skin grafting, local skin flaps, pedicles flaps, and up to utilization of a free flap for soft tissue coverage. More recently, we are employing so-called hybrid reconstruction (Figure 2), whereby we are utilizing regenerative medicine modalities in the management of composite tissue loss. These include the use of dermal regenerative templates such as Integra (Integra LifeSciences, Plainsboro, NJ), external tissue expanders such as Dermaclose (Wound Care Technology, Chanhassen, MN), extra-cellular matrices such as ACell, local rotation flaps, and free tissue transfer to achieve a functional residual limb. This also entails proceeding directly to free flap, if necessary, as soon as wounds allow, skipping many or all lower rungs of the traditional reconstructive ladder (Figure 3). Regenerative medicine therapies offer significant enhancements to traditional coverage methods, such as extending the indications for a particular rung on the ladder.

In addition to regenerative medicine modalities, we've additionally incorporated nanotechnology into amputation wound management. We frequently employ dressings which release nanocrystalline silver crystals which can eradicate bacteria.<sup>12</sup> Studies suggest that this technology alters wound inflammatory events and facilitates the early phase of wound healing.<sup>13</sup> We strive for amputated limb length salvage for today's Wounded Warriors with the end-state in mind of enhancing their performance 30 to 40 years from now, when a 25% to 65% increase in energy expenditure above baseline means a mobile versus wheelchair-dependent 60-year-old.

We use an abundant amount of Integra (originally designed for burn management), which is a bilayer matrix wound dressing consisting of cross-linked bovine tendon collagen and glycosaminoglycan with a semi-permeable silicone layer that controls water vapor loss, provides a flexible adherent covering, allows increased shear strength, and appears to biodegrade for cell and capillary ingrowth.<sup>14</sup> The dressing is typically



**Figure 2.** The hybrid reconstructive ladder employs techniques such as dermal regenerative templates, external tissue expanders, extra-cellular matrices, and nerve conduits to achieve the most functional residual limb.



**Figure 3.** The hybrid reconstructive ladder express. The complex wound care demanded by today's high energy amputations may require skipping some or all lower rungs of the traditional reconstructive ladder and proceeding directly to the highest levels of limb salvage, including pedicle or free flaps combined with newer regenerative medicine therapies.

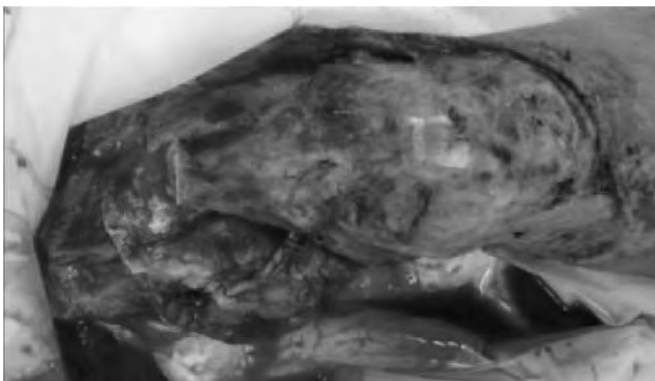
utilized in conjunction with negative pressure wound therapy (NPWT) and is followed by an ultra-thin STSG (12 micrometers) at approximately two to three weeks. The dermal substitute effectively recreates adequate subcutaneous tissue to allow for gliding or movement of the skin over the muscle and it has been objectively demonstrated that the elastic properties of areas treated with Integra are comparable to normal skin.<sup>15</sup>

There have been several case reports describing the use of NPWT to granulate areas of soft tissue loss such as over tendons and fascia. An adjunct to NPWT that we have employed is the use of porcine urinary bladder matrix as a biologic scaffold. This biologic scaffold attracts adult stem cells<sup>16</sup> to the site of injury and converts them into active progenitor cells.<sup>17</sup> It also sets up non-crosslinked temporary scaffolding for tissue reconstruction, mimicking surrounding healthy tissue.<sup>18</sup> This scaffolding maintains an intact epithelial basement membrane<sup>19</sup> and typically results in a robust bed of granulation tissue. STSG would have likely failed if applied directly to the retinaculum and patella ligament (Figure 4). In this case, 100% take of the STSG was achieved.

Several authors have described the use of free or rotational flaps to facilitate lower extremity amputation length salvage.<sup>20-23</sup> Although the latissimus free flap has been the most frequently reported, other microvascular techniques have included use of an anterolateral thigh fasciocutaneous flap<sup>24</sup> and parascapular flaps for transtibial amputation salvage. Kasabian, et al. published the largest series with 22 patients and were able to maintain a below-knee amputation in all patients.<sup>20</sup>

## CONCLUSION

The preservation of residual limb length in lower extremity amputations is crucial to optimize prosthetic fitting for the amputee to allow for his or her recovering maximal function with ambulation. Applying principles of the reconstructive ladder may facilitate the achievement of soft tissue coverage, which is paramount in lower extremity amputees.



**Figure 4.** This patient, with significant soft tissue loss above a high below-knee amputation, underwent treatment with ACell followed by split thickness skin grafting.

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## Heterotopic Ossification Resection after Open Peri-articular Combat-Related Elbow Fractures

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### INTRODUCTION

Heterotopic ossification (HO) can have devastating effects on achieving a successful outcome in the treatment of elbow trauma, resulting in prolonged rehabilitation and subsequent procedures to improve range of motion (ROM) lost to bone impingement and arthrofibrosis.<sup>1</sup> Surgical excision of heterotopic ossification to improve functional elbow ROM has been shown to be safe and effective among several published series of closed, civilian trauma.<sup>2-9</sup> Given that combat-related injuries are characterized by high energy blast and ballistic mechanisms, wide zones of injury, soft tissue loss, systemic inflammation from concomitant injuries, and high incidence of traumatic brain injury (TBI), it can be expected that combat-related peri-articular elbow injuries may represent the worst case scenario for HO development and elbow injury outcomes. Surgical treatment of elbow contracture secondary to heterotopic ossification in blast-injured elbows has not yet been evaluated in the literature, to our knowledge. The objective of this analysis is to review the outcomes of HO resection to improve ROM in combat-related, open, peri-articular elbow fractures complicated by heterotopic ossification.

### METHODS

A retrospective review of open, peri-articular, combat-related elbow fractures treated at the National Naval Medical Center at Bethesda and Walter Reed Army Medical Center (now Walter Reed National Military Medical Center) was approved by the respective institutional review boards. Open, peri-articular elbow injuries were defined as open fracture within 5 cm of the radiocapitellar or ulnohumeral joints. Patients treated with upper extremity amputation within the first 24 hours of injury were excluded. A review of our institution's electronic surgical scheduling system revealed surgical treatment of 132 consecutive peri-articular injuries to the elbow in 128 combat-injured male patients treated at our facility between 2004 and 2011. Heterotopic ossification was defined as ectopic bone formation within the zone of injury after initial definitive management. Ninety-two (69%) elbows developed HO on radiographic follow-up. Inclusion criteria were patients who underwent primary elbow HO excision surgery with concurrent capsulectomy/lysis of adhesions to improve flexion-extension arc of motion. Of the 43 elbows on which HO excision was performed, 13 elbows were excluded from

review: seven had inadequate pre- and post-operative ROM documentation, two had proximal forearm amputations requiring HO excision for pain or prosthetic fitting, and three had procedures primarily to improve pronation-supination motion.

Abstracted data was collected from inpatient and outpatient electronic medical records. Radiographs demonstrating the most severe HO were used to determine Hastings classification.<sup>10</sup> Outcomes were derived from electronic medical record capture of follow-up orthopaedic

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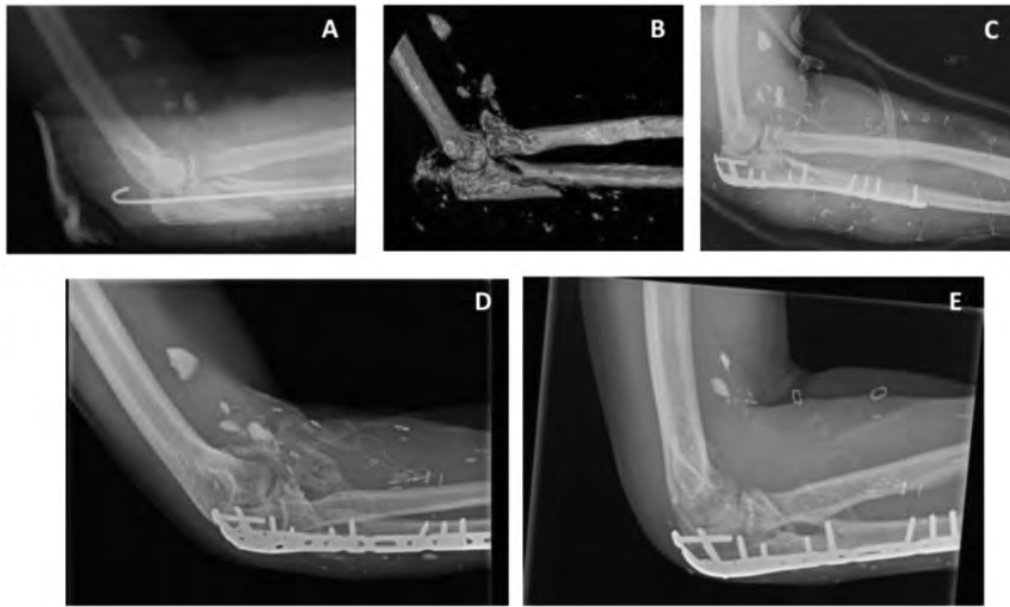


LCDR(s) Wilson earned a bachelor of science degree in biology from Gettysburg College in 2003. Following graduation, he accepted the Armed Forces Health Profession Scholarship and was commissioned into the U.S. Navy. He earned his doctor of medicine degree at Jefferson Medical College in Philadelphia, PA. Lcdr(s) Wilson completed his orthopaedic surgery internship at the National Naval Medical Center in 2008 and was selected for orthopaedic surgery residency training. During his research fellowship, he focused his work on a variety of projects related to spine biomechanics, degenerative disc disease, hip arthroscopy, and outcomes of combat extremity injuries. He authored a protocol that was awarded \$205,000 in funding from the Navy Bureau of Medicine and Surgery Clinical Investigations Program. After graduation in June 2013, Lcdr(s) Wilson will begin his orthopaedic surgery career at Naval Hospital Camp Lejeune, North Carolina. His future plans include a fellowship and career in adult reconstructive surgery.

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**Figure 1.** Serial radiographs of treatment course. Twenty-seven-year-old active duty male sustained improvised explosive device blast injury to the left upper extremity while serving in Afghanistan as an explosive ordinance disposal team leader, resulting in a Gustilo-Anderson type IIIB open proximal ulna and radial head fracture-dislocation. He was treated for TBI. He underwent irrigation and debridement and provisional stabilization prior to arrival at our facility. Radiographs (A) and CT 3D reconstruction (B) are shown. He underwent serial irrigation and debridement. After antero-lateral thigh free tissue transfer, we performed open reduction internal fixation (C) 11 days after injury. After three postoperative months, he returned with elbow HO and contracture (D) that was refractory to therapy and splinting. His arc of motion was 25 to 65 degrees of flexion. HO resection was performed six months after definitive fixation, followed by intensive occupational therapy. Post-resection (two weeks postoperative) radiographs are shown (E). At two years follow-up, his arc of motion was 30 to 115 degrees of flexion.

clinic and occupational therapy encounters, which routinely include ROM and complication details. Serial radiographs of a case example are displayed in Figure 1.

### Surgical Details and Rehabilitation

A full discussion of surgical techniques and rehabilitation principles is beyond the scope of this report. After review of operative reports, the approach utilized was dictated by the anatomic location of the focus of heterotopic bone, presence of flaps, and need for hardware removal. HO prophylaxis regimens varied throughout the collection period. Routine follow-ups with exam occurred generally at 2 weeks, 6 weeks, 12 weeks, 6 months, and annually until loss of follow-up, after separation from the active duty military to the Veterans Affairs system.

The ROM at final follow-up, including cases of failed and repeat procedures, was used for outcome calculations, as per the intent-to-treat principle. Statistical analysis of the data included basic summary calculations for demographic variables and testing of the pre-operative and post-operative ROM between groups with independent samples *t*-tests, equal variances not assumed.

## RESULTS

The demographic and injury characteristics for the cohort are summarized in Table 1. The mean age was 25 years (range, 19 to 42

years), and mean follow-up was 35 months (range, 12 to 51 months). The mean time from initial definitive management until HO resection was 10 months, (range, 3 to 18 months). The mechanism of injury was most commonly explosive blasts (90%, 27/30). The majority of injuries were Gustilo-Anderson type III injuries. Ten elbows (33%) were IIIB, requiring pedicled or free tissue transfer for definitive wound closure. Four (13%) were IIIC, associated with vascular injury requiring repair or revascularization procedure at time of injury. Seventeen elbows (57%) had associated nerve injuries. Fifty-three percent of patients (16/30) were diagnosed with TBI after known head injury or after mandatory TBI screening (Table 1).

### Analysis of HO Resection Outcomes

The mean preoperative flexion-extension arc of motion was 36.4 degrees, compared with mean postoperative arc of motion of 83.6 degrees. A mean sustained gain of 47.2 degrees of flexion-extension ROM (range, -15 to 110 degrees, Table 2) was seen across the sample. The arc of motion preoperatively and postoperatively for the injured elbows in this series is significantly narrowed from the normal range of motion (0 to 140 degrees) for uninjured elbows (Figure 2). No significant difference was observed in the ROM gains between elbows with and without TBI diagnosis ( $p = 0.102$ ), a well-established risk factor for heterotopic ossification.

**Table 1. Demographics and Injury Characteristics**

Age	25	(19-42) years
Follow-up	35	(12-51) months
Mechanism	27	(90%) blasts
	3	(10%) gun shot wounds
TBI	16	(53%) of patients
Time to HO resection	10	(3-18) months

**Fracture**

Location	# elbows	percentage
Humerus	20	67%
Ulna	23	77%
Radius	7	23%
Combined	13	43%

**Prior Procedures (mean)**

Downrange	2.7	procedures
Debridements	5.2	procedures

**Gustilo-Anderson**

Type	# elbows	percentage
I	0	0%
II	8	27%
IIIa	8	27%
IIIb	10	50%
IIIc	4	20%

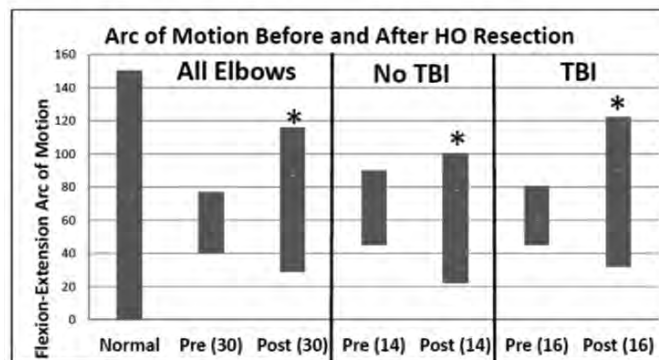
**HO Classification**

Class	# elbows	percentage
I	0	0%
IIa	10	33%
IIb	5	17%
IIc	14	47%
III	1	3%

**Table 2. Degrees of Motion Before and After HO Resection**

Variable	#	Before	After	Gain	p value
All elbows	30	36.4	83.6	47.2	< 0.001
TBI	16	38.3	78.1	39.8	< 0.001
No TBI	14	34.2	89.9	55.7	< 0.001
Flap	10	37.9	79.6	41.7	0.003
No flap	20	35.6	85.6	50.0	< 0.001
Nerve injury	17	39.2	78.4	39.2	< 0.001
No nerve injury	13	32.6	90.4	57.8	< 0.001

The mean degrees of motion pre-operative, post-operative, and total gain measurements is displayed for the entire cohort (all elbows), and for elbows associated with and without TBI, need for flap coverage, and presence of nerve injury. Analysis of all subsets across the cohort demonstrated a significant gain in motion.



**Figure 2.** Arc of motion before and after HO resection. Arc of motion is displayed for normal elbows, pre-operative elbows, post-operative elbows for entire cohort, elbows without concomitant TBI, and elbows in individuals with TBI diagnosis. \*denotes a significant difference between pre- and post-operative ROM values.

We noted significant mean gains in ROM from pre-operative measurements to final post-operative measurements across the sample, including subsets with and without potential confounding factors (Table 2). Sub-analysis of only elbows with concomitant TBI, need for flap, and nerve injury continued to demonstrate a significant gain in ROM in each subset, with and without the potential complicating variable. We found no significant difference in the pre-operative and post-operative motion for elbows with or without each of the potential complicating variables (Table 3). While there appeared to be a trend for elbows without TBI (55.6 vs. 39.9 degrees), flaps (50.0 vs 41.7 degrees), or nerve injuries (57.8 vs 39.2 degrees) to achieve a greater gain in range of motion, no statistically significant difference was found.

**Table 3. Comparison of Pre- and Post-Operative Motion by Potential Risk Factors**

Variable	Before	p value	After	p value	Gain	p value
TBI	38.3	0.358	78.1	0.115	39.9	0.102
No TBI	34.2		89.9		55.6	
Flap	37.9	0.419	79.6	0.300	41.7	0.274
No flap	35.6		85.6		50.0	
Nerve injury	39.2	0.538	78.4	0.203	39.2	0.136
No nerve injury	32.6		90.4		57.8	

A student's *t*-test of means was performed for the pre-operative, post-operative, and total gain measurements for elbows with concomitant TBI, need for flap coverage, and presence of nerve injury. No statistically significant differences were found among the mean pre-operative, post-operative, or gain measurements among elbows with any of the potential risk factors for stiffness.

## Complications

HO resections were complicated by one intraoperative distal humerus fracture, six episodes of recurrent arthrofibrosis (less than 15 degree improvement), and one re-injury of a previously-injured posterior interosseous nerve (PIN). Four of the six recurrent arthrofibrosis cases went on to repeat lysis of adhesion and manipulation under anesthesia procedures, with no significant lasting improvement. The PIN palsy improved with observation.

## DISCUSSION

The current study may represent the largest series of HO resections involving peripheral nerve injuries, open fractures, or wounds requiring flap coverage. It can be implied that nerve injury and soft tissue deficits represent more serious insults to the peri-articular soft tissues that are involved in the elbow contractures, however, these characteristics were not found to play a role in the ROM outcomes of surgical resection of HO. The results of the present series demonstrate generally comparable results to HO resection following lower energy, closed injuries, with slightly decreased ROM gains and similar complication rates.

Resection of HO to improve functional elbow ROM has been shown to be successful in varied clinical settings, with documented improvements in objective performance scores and increases in ROM from 49 to 94 degrees.<sup>2-8,11</sup> Ring and Jupiter reported the successful operative release of complete ankylosis in 20 elbows among 15 severe trauma and burn patients, with a mean sustained arc of motion of 81 degrees for the burn cohort and 94 degrees for the trauma cohort.<sup>11</sup> The soft tissue injuries described are less severe than the combat injuries in the current series. Also, none of the Ring and Jupiter patients had concurrent TBI.<sup>11</sup> More recently, Baldwin, et al.<sup>3</sup> presented a mean gain in arc of motion of 49 degrees in a cohort of patients with TBI, direct elbow trauma, and combined etiologies. Our mean increase in flexion-extension arc of motion of 47.2 degrees is comparable, considering the complexity of these types of patients. Similarly, Baldwin, et al. reported that timing of resection and neurologic characteristics had no effect on outcomes.<sup>3</sup>

Our rate of complications (26.7%), including recurrence, is similar to the rates reported in other series.<sup>2-9,11</sup> Ring and Jupiter reported clinically significant recurrent HO in one-third of their trauma cohort.<sup>11</sup> Means for identifying those limbs at risk for complication or recurrence have not been established. Given the unremarkable influence of potential risk factors for HO and contracture in this and other series, potential etiologies for recurrent contracture remain unclear.

The shortcomings of this analysis are related to its retrospective nature and multiple treatment regimens carried out by multiple surgeons. No consistent regimen of chemo- or radiotherapy HO prophylaxis was used; however, recommendations for safe prophylaxis regimens after elbow injury or surgery have not yet been established in other patient populations.<sup>1,2,9</sup>

Based on our observations, HO resections in combat-related blast injuries to the elbow can produce reliable gains in the arc of elbow motion. Blast injuries are not commonly encountered in the orthopaedic community; nevertheless, it may be noteworthy that the utility of HO resections with concurrent lysis of adhesions is maintained following these complex injuries, even in what may be the worst-case scenario for post-traumatic elbow HO. Elbow contractures associated with open fracture, soft tissue loss, and nerve injury should not preclude attempts at improving ROM with surgical excision of HO.

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An Innovative Culture System for the Study of Post-Traumatic Heterotopic Ossification

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INTRODUCTION

Following orthopaedic trauma, injured muscle tissue becomes populated with mesenchymal progenitor cells (MPCs) containing a cell surface epitope profile and multidifferentiation potential that is characteristic of bone marrow-derived mesenchymal stem cells (MSCs).<sup>1</sup> Our laboratory has hypothesized that these cells, which appear to be intimately involved in the healing process, also play a key role in wound healing pathologies such as heterotopic ossification (HO).

HO is defined as aberrant mature bone formation in non-osseous tissue. During recent military operations, there has been a significant increase in the prevalence of heterotopic ossification in the combat-wounded patient population, with rates as high 64%.<sup>2</sup> Symptomatic HO can significantly worsen morbidity, complicating internal fixation of fractures and prosthetic fitting in amputees, and requiring additional surgical procedures for symptomatic HO excision. Understanding the pathogenesis of this condition is essential in the prevention and potential treatment of the disease.

We hypothesize the physical microstructure (fibrosis) within a wound dysregulates muscle regeneration, working synergistically with inflammatory factors to confuse local MPC populations and generate an early osteoinductive region. In this study, our specific aims were (1) to examine the structure of decellularized early-stage traumatized muscle tissue taken from patients that develop HO and identify non-cellular structural components that may be instrumental in pathological wound development, (2) to create a biomimetic in vitro culture platform replicating the wound microarchitecture and evaluate the osteogenic potential of this microarchitecture compared to conventional 2D cell culture approaches. Our overall goal is to create an in vitro model system for studying the development of HO.

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MATERIALS AND METHODS

Tissue Decellularization

Surgical waste muscle tissue was obtained from consented patients injured during Operation Enduring Freedom; these patients were radiographically observed as developing HO in subsequent months (Figure 1). Tissue was sectioned into 5 mm<sup>3</sup> segments and incubated overnight in 0.1% sodium dodecyl sulfate (SDS) surfactant solution to remove all cellular and lipid components.

Scanning Electron Microscopy (SEM)

Decellularized tissue samples were fixed with paraformaldehyde/glutaraldehyde, secondarily fixed with osmium tetroxide, serially diluted with ethanol, hexamethyldisilazane (HMDS) critically point dried and sputter-coated with gold for SEM imaging.

Collagen Nanofibers

Collagen nanofibers were electrospun to replicate the tissue microarchitecture in vitro. Lyophilized calf-skin collagen I was dissolved in hexafluoroisopropanol (HFIP) at 3% w/w and electrospun onto grounded aluminum plates.<sup>3</sup> The fiber matrix was manipulated into coverslips (for 24-well plates samples) and 15-cm culture dishes for



**Figure 1.** Radiographs of a soldier injured during Operation Enduring Freedom, showing the development of heterotopic ossification (HO) in the weeks following blast injury.



large-scale cell culture. Samples were glutaraldehyde vapor-saturated for 15 minutes for crosslinking stability, and thoroughly rinsed with phosphate buffered saline (PBS).

### Cell Isolation and Culture

MPCs were harvested using a previously described protocol.<sup>1</sup> Tissue identical to that used for decellularization was minced, incubated in collagenase II, and filtered through a 40- $\mu$ m cell strainer to obtain a cell solution. The cell solution was incubated for two hours on tissue culture-treated plastic flasks and non-adherent cells were vigorously rinsed away to obtain a mesenchymal progenitor population. Cells were expanded in high-glucose DMEM culture media containing 10% fetal bovine serum. To evaluate the osteogenic potential of the traumatized-tissue-isolated MPCs, cells were expanded on collagen-coated and fiber matrix dishes for 3 weeks, before ensuing culture in osteogenic media (Invitrogen) for 2 weeks. Calcium deposition was measured with alizarin red, and osteoblast activity was assayed with alkaline phosphatase. Protein lysates were also collected for western blotting analysis.

### RESULTS

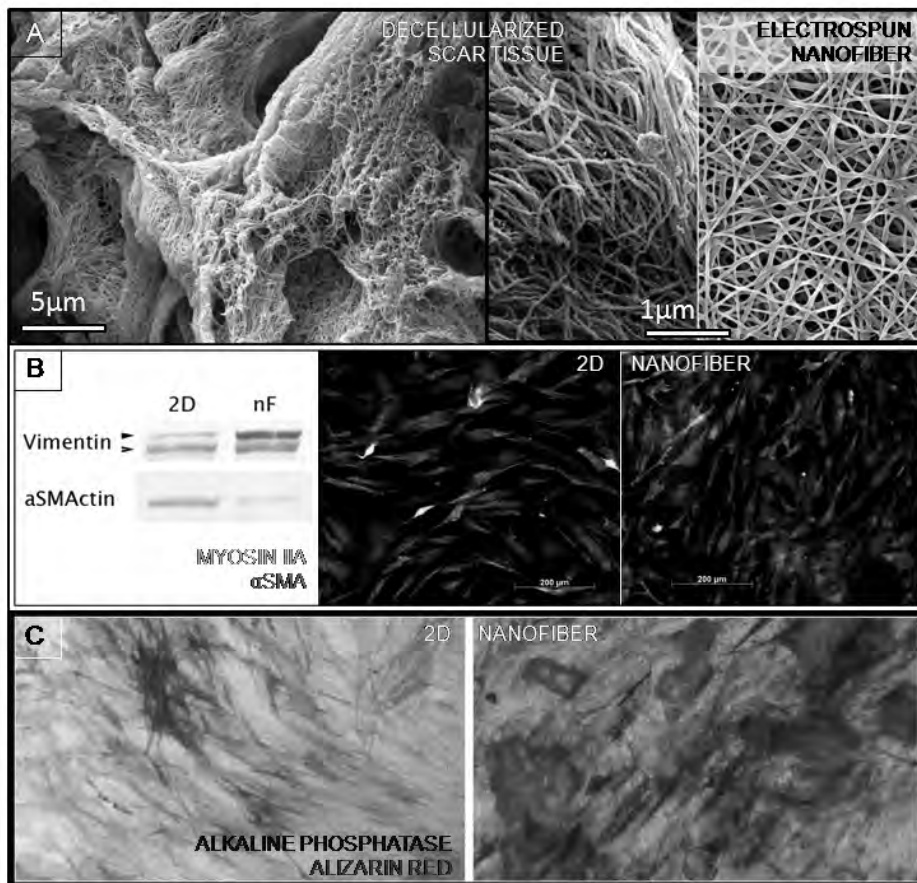
We have evaluated the physical microenvironment of traumatized muscle tissue in patients radiographically confirmed to form HO by

lysing all cellular components from the tissue, leaving a preponderance of nanofiber matrix. This nanofiber component was identically replicated with electrospun collagen scaffolds (Figure 2A). MPCs cultured on this fiber surface have a significantly increased osteogenic potential, with greatly increased calcium deposition (as measured by alizarin red) and alkaline phosphatase expression, compared to cells cultured on 2D collagen (Figure 2C).

Western blotting analysis revealed increased vimentin expression for MPCs cultured on fiber substrates prior to osteogenesis (Figure 2B), as well as a decrease in alpha smooth muscle actin ( $\alpha$ SMA), a prominent fibroblast marker. Vimentin has been shown to be essential for cartilage formation,<sup>4</sup> which is hypothesized to precede endochondral ossification of HO. It has also been shown that decreases in vimentin correlate with osteoblast differentiation and bone formation, perhaps acting as a trigger for osteocalcin signaling.<sup>5</sup>

### DISCUSSION

Fibrotic tissue that forms in traumatized muscle (i.e., scar) is known to precede HO, a debilitating formation of ectopic bone in soft tissues whose cellular mechanisms are poorly understood. SEM characterization of fibrotic muscle tissue reveals an abundant fibrous network, a component which our findings indicate may be sufficient to promote cellular osteogenic differentiation in vitro. Recent studies show



**Figure 2.** (A) Scanning electron microscopy (SEM) images of traumatized muscle tissue, decellularized with sodium dodecyl sulfate (SDS) detergent. The abundant nanofiber matrix was recreated with electrospinning. (B) Mesenchymal progenitor cells (MPCs) isolated from the traumatized muscle were cultured on 2D collagen and nanofiber substrates. Western blotting and immunofluorescence microscopy reveal an increase in vimentin (intermediate filament, known to play key role in chondrogenesis, fibrosis, and osteogenesis) and a decrease in alpha smooth muscle actin (fibrosis marker) for MPCs cultured on fiber matrix. (C) MPCs cultured on nanofiber matrix in osteogenic media have a robust increase in alizarin red staining (calcium excretion) and ALP expression, indicating a heightened sensitivity to osteogenic induction.

that a collagen fiber matrix precedes bone formation during development, indicating bone may spontaneously form if collagen matrix is present, likely from cells secreting mineral matrix through exocytosis.<sup>6</sup> This developmental cascade is replicated in our experimental results, which show a robust increase in calcium secretion from MPCs cultured on a biomimetic nanofiber matrix. The increase in vimentin expression before osteogenic induction correlates well with literature, with a corresponding decrease in fibrotic markers ( $\alpha$ SMA) indicating a shift into preosteogenic state. The sensitivity of osteogenic phenotype to the nanofiber matrix suggests that it is likely a key component for properly elucidating HO development using in vitro cellular models.

This study helps to recreate in vivo elements associated with HO in a representative culture system, providing a suitable platform for studying the cellular parameters responsible for such pathological wound healing. Surface topography appears to contribute substantially to osteoinductive wound healing, making it an essential component for investigating and minimizing the risk of HO.

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## Outcomes Following Cervical Disc Arthroplasty: a Retrospective Review

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### INTRODUCTION

Cervical radiculopathy is a common diagnosis in both civilian and military populations that causes significant disability and loss of work.<sup>1-3</sup> In the military population specifically, radiculopathy may result in an inability to perform soldier tasks proficiently, preventing deployment and decreasing force strength. Patients may present with a constellation of symptoms, including dermatomal radicular pain and paresthesias, weakness, and sometimes myelopathic findings such as hyperreflexia and positive rudimentary reflexes (Hoffman's, Babinski, etc.).<sup>4-7</sup> Several surgical options are available for the treatment of cervical radiculopathy following a trial of conservative management. These options include posterior decompression and fusion, anterior discectomy and fusion, and cervical disc arthroplasty (CDA). CDA has been espoused in the literature as a viable alternative to anterior cervical discectomy and fusion (ACDF), with the added theoretical benefit of preventing adjacent segment degeneration, though this remains to be seen.<sup>8-12</sup> Based on our increasing experience in performing CDA, we sought to determine the outcomes of all patients who underwent this procedure at our institution.

### MATERIALS AND METHODS

Following approval from our institutional review board, the surgical database at this institution was queried to identify all patients who had undergone CDA between August 2008 and August 2012. This search yielded 316 total patients. Of the total, 34 were lost to follow-up, leaving 282 patients for review. All construct types (single level CDA, hybrid, and multilevel CDA) were included. All data were collected via a retrospective chart analysis, which included inpatient and outpatient clinical notes, surgical databases, and radiographs. Data collected include patient demographic information (age, sex, tobacco use, body mass index (BMI)), patient-centered outcomes (relief of pre-operative symptoms, incidence/resolution of axial neck pain in the post-operative period, return to full activity), complications (recurrent laryngeal nerve injury, dysphagia, post-operative respiratory compromise, esophageal/tracheal disruption), and radiographic parameters (increase in disc height, segment range of motion, evidence of loosening, migration, or subsidence).

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### RESULTS

There were 219 males (77.7%) and 63 females (22.3%). The average length of follow-up was 11.2 ( $\pm$  11.0) months. The average patient age was 42.1 ( $\pm$  8.4) years. The average BMI was 27.8 ( $\pm$  3.7) kg/m<sup>2</sup>. The most common levels addressed at the time of surgery were C5-6 and C6-7 (56.7% and 58.9%, respectively). The next most frequently diseased levels were C4-5 (19.5%), C3-4 (8.2%), and C7-T1 (2.5%). A majority of patients underwent single level CDA (59.9%), and 22.3% underwent two-level hybrid (CDA/ACDF) constructs. Twenty-five patients (8.9%) underwent two-level CDA; 11 (3.9%) of those patients had contiguous levels addressed, and 14 (5.0%) had non-contiguous levels addressed. Three patients had three-level contiguous CDA (Table 1). Of the patients studied, 73.8% (209) were active duty at the time of surgery. The Prestige™ (Medtronic, Memphis, TN) cervical arthroplasty system was utilized in the majority of patients (94.5%), while the ProDisc-C system (Depuy Synthes, Paoli, PA) was utilized in the remainder of patients (5.5%).

The primary indications for surgery were radiculopathy (86.5%), myelo-radiculopathy (9.2%), and myelopathy (1.8%). Flexion and extension lateral radiographs were available in 199 patients (60.9%), and the mean range of motion (ROM) at the CDA levels was 7.5 ( $\pm$  4.1) degrees at latest follow-up. Of the 282 patients reviewed, 250 (88.7%) experienced complete relief of pre-operative complaints. Twenty-six patients (9.2%) had incomplete relief of their symptoms, to include complaints of persistent axial neck pain, residual paresthesias, or radiculopathy. Some patients (2.1%) experienced initial relief of symptoms, but later developed recurrence during follow-up. We found 52 (18.4%) patients who complained of axial neck pain at three months or greater, but that eight (15.3%) had resolution of their pain at one-year follow-up with conservative therapy, and 11 (21.2%) stated that their pain did not require treatment or affect their return to activity. A vast majority of patients were able to return to full activity (92.2%). In the active duty population, there was a 90.4% rate of return to full activity. Of the 24 patients who did not return to full activity, 20 were active duty at the time of surgery and required a medical board for inability to perform their job in the necessary capacity (Table 1).

The complication rate was 14.5%, which included dysphagia (8.9%), recurrent laryngeal nerve injury (3.2%), post-operative hematoma (0.4%), nerve root injury (0.4%), spinal cord injury (0.4%), and superficial infection (0.4%). There was a 5.3% re-operation rate, including conversion of CDA to ACDF, revision of ACDF or CDA, posterior decompressions, adjacent segment degeneration, and hematoma decompression (Table 2).

When the data were analyzed based on construct type, we found similar outcomes between single level, two-level hybrids, and two- and three-level CDA (Table 3). The multi-level hybrid constructs (most commonly CDA/ACDF/CDA) showed a decreased rate of symptom relief and return to full activity (77.8% and 72.2%, respectively), and higher rates of persistent axial neck pain (50%) and dysphagia (22.2%) (Table 3).

**Table 1. Patient Information and Outcomes**

Total patients	282
Men	71.7% (219)
Women	22.3% (63)
Age	42.1 ± 8.4 years
Body mass index	27.8 (± 3.7) kg/m <sup>2</sup>
Tobacco use	26.6% (75)
Active duty military	73.8% (208)
Revision surgery	5.7% (16)
Average follow-up	11.2 ± 11.0 months
Levels of disease	
C3-4	8.2% (23)
C4-5	19.5% (55)
C5-6	56.7% (160)
C6-7	58.9% (166)
C7-T1	2.5% (7)
Primary indication for surgery	
Myelopathy	1.8% (5)
Radiculopathy	86.5% (244)
Myelo-radiculopathy	9.2% (26)
Neck pain	0.7% (2)
Outcomes	
Relief of neurologic symptoms	92.9% (262)
Complete relief of symptoms	88.7% (250)
Persistent axial neck pain (>3 mos)	18.4% (52)
Return to full activity	92.2% (260)
Return to active duty military	90.4% (189/209)

**Table 2. Adverse Outcomes**

Complications	
Recurrent laryngeal nerve injury	3.2% (9)
Dysphagia	8.9% (25)
Post-op hematoma	0.4% (1)
Infection	0.4% (1, superficial)
Spinal cord injury	0.4% (1)
Dural tear	0.4% (1)
Recurrence of neurologic symptoms	2.1% (6)
Reoperation during follow-up	5.3% (15)
Posterior decompression	2.1% (6)
Adjacent level	1.8% (5)
Conversion to anterior cervical discectomy and fusion	0.7% (2)
Hematoma decompression	0.4% (1)

**Table 3. Outcomes Based on Construct Type**

Single level CDA	59.9% (169)
Complete symptom relief	90.1%
Return to full activity	93%
Persistent axial neck pain	15.8%
Dysphagia	5.8%
Recurrent laryngeal nerve injury	2.9%
Two-level hybrid	22.3% (63)
Complete symptom relief	90.6%
Return to full activity	90.6%
Persistent axial neck pain	12.5%
Dysphagia	12.5%
Recurrent laryngeal nerve injury	3.1%
Two-level contiguous CDA	3.9% (11)
Complete symptom relief	96.0%
Return to full activity	96.0%
Persistent axial neck pain	24%
Dysphagia	8%
Recurrent laryngeal nerve injury	4%
Three-level hybrid	6.4% (18)
Complete symptom relief	77.8%
Return to full activity	72.2%
Persistent axial neck pain	50%
Dysphagia	22.2%
Recurrent laryngeal nerve injury	0.0%

CDA, cervical disc arthroplasty

## DISCUSSION

To our knowledge, this is the largest, non-funded, single center review of CDA. When analyzing all construct types that included CDA at our institution, we found an 88.7% rate of complete pre-operative symptom relief and a 92.2% rate of return to full activity, with maintenance of 7.5 degrees at each CDA level on average. This included a majority subset population of active duty service members, which demonstrated a 90.4% rate of return to active duty. Furthermore, our review demonstrates a low complication rate with regard to post-operative dysphagia (8.9%) and recurrent laryngeal nerve injury (3.2%).

The rate of dysphagia in this review (8.9%) is lower than that reported in the literature (28% to 57%),<sup>13-17</sup> which is most likely due to the fact that dysphagia is often underreported clinically.<sup>14</sup> Most surgeons at our institution counsel patients to expect a normal two- to three-month period of “settling” posterior neck pain postoperatively. However, we found a high rate (18.4%) of persistent neck pain in the post-operative period, which was defined as pain lasting longer than three months and/or neck pain that required further non-operative intervention (extended physical therapy, facet injections, or nerve ablation). Unfortunately, we were unable to correlate this rate of axial neck pain with the presence

of pre-operative axial neck pain as a presenting symptom, as this was unreliably documented.

Another pertinent finding was the lower rate of pre-operative symptom relief and return to full activity following multilevel hybrid constructs (77.8% and 72.2%, respectively) and a higher rate of dysphagia and persistent post-operative axial neck pain. These patients likely experienced a higher severity of pre-operative symptoms compared to the one- and two-level groups, with 27.8% of patients in the multi-level hybrid group experiencing myelopathic complaints compared to 10% in the remainder of constructs. These larger constructs also likely required longer operative times, which may have led to the increased rates of dysphagia.

While ACDF continues to be the “gold standard” in the management of cervical degeneration,<sup>18-21</sup> several recent studies have shown comparable short- and mid-range follow-up outcomes with CDA.<sup>8-12,22,23</sup> Long-term ACDF studies have shown approximately 3% per year adjacent segment degeneration rate.<sup>24</sup> In a mid-term follow-up study, Sun, et al. found a significantly decreased rate of adjacent segment degeneration in patients who had undergone CDA compared to ACDF (17.6% versus 60.4%, respectively) with five-year follow-up.<sup>25</sup> While these intermediate results are promising, there are currently no long-term studies (> 10 years) to demonstrate reduction in adjacent segment degeneration compared to ACDF.

Cervical disc arthroplasty is increasing in popularity for the treatment of cervical radiculopathy and myelopathy. To our knowledge, this is the largest, non-funded, single center study on this topic. Our review found an 89% rate of complete pre-operative symptom relief, with greater than 90% of patients returning to their pre-operative levels of activity and an acceptably low rate of complications. While there are currently no long-term studies demonstrating the superiority of CDA over ACDF, short- and mid-term results have shown that CDA is a safe and reliable alternative to ACDF for properly selected patients. However, further studies are necessary to determine if there is a long-term reduction in adjacent segment degeneration or re-operation.

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## The Use of Dilute Dakin's Solution for the Treatment of Invasive Fungal Infection in the Combat Wounded: a Case Series

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### INTRODUCTION

**B**last injury-related invasive fungal infections (IFIs) have dramatically shaped the treatment and outcomes of individuals injured in current combat operations. These infections are traditionally seen in immunocompromised patients in the civilian literature and are typically managed with aggressive debridement and systemic antifungal medications, with limited success.<sup>1-8</sup>

In the current report, we describe the use of dilute Dakin's solution (0.025% buffered sodium hypochlorite solution)<sup>9</sup> and negative pressure wound therapy (NPWT) as an adjunct to the aggressive surgical debridement required to address aggressive fungal infections. Sodium hypochlorite solution has been previously utilized for treatment of combat-related wound infections since the Civil War;<sup>10</sup> however, the use of dilute Dakin's solution has not been described to date in the management of IFI in patient with severe traumatic wounds.

### CASE STUDY

Our patient is a 24-year-old male Marine who sustained injuries while conducting combat operations in support of Operation Enduring Freedom. His vehicle encountered an improvised explosive device (IED) causing multiple injuries, including a left transfemoral amputation, right transtibial amputation, and transradial amputation; additionally, he sustained a severely mangled left upper extremity. His initial management included obtaining hemorrhage control, decontamination, and resuscitation in Afghanistan. He underwent a revascularization utilizing a local vein graft for his left upper extremity. Once stabilized, he was evacuated to Landstuhl Regional Medical Center and underwent additional irrigation and debridement procedures prior to transferring to our facility. Upon arrival at our facility six days from the date of injury, he was febrile to 104°F with exposed left forearm vein grafts, an open abdomen, and malfunctioning Wound Vacuum Assisted Closure (V.A.C.)<sup>TM</sup> (KCI, San Antonio, TX). Therefore, a decision was made to urgently take him to the operating room for wound assessment and aggressive debridement and irrigation.

Intraoperatively, his right knee disarticulation appeared viable. However, his vein grafts were nonviable and appeared necrotic. He did have a dopplable arch, which is presumed to be from dorsal collateral flow. His right transradial amputation was largely nonviable and the radius and ulna were removed, along with large amounts necrotic soft tissue. The high left transfemoral amputation was malodorous and had obvious necrosis (Figures 1 to 3). Following aggressive debridement and irrigation of all wounds, the patient was started on broad spectrum

antibiotics and tissue was sent for culture. Based on the appearance and presence of extensive necrosis in all wounds, a decision was made to return to the operating room in 24 hours for repeat debridement and irrigation. During the next procedure, due to the extent of tissue necrosis around the left transfemoral amputation, his left transfemoral amputation was revised to a hip disarticulation. Several specimens from the left lower extremity were sent to pathology for histological evaluation, as well as to microbiology. Histologic analysis revealed angioinvasive fungal elements. Wound cultures eventually grew *Aspergillus fumigatus*.

On post-injury day 10, the patient underwent repeat debridement of his bilateral lower extremity wounds. During this trip to the operating room, he also required a left transradial amputation, as a result of extensive tissue necrosis, and an inability to maintain adequate tissue perfusion to the hand. Serial debridements continued every 48 hours. Once positive fungal cultures were obtained, Dakin's solution was added to the intra-operative irrigation solution at the next debridement. The patient's left hip disarticulation wound was then treated with Dakin's solution infused utilizing Instill therapy Wound V.A.C.<sup>TM</sup>. This allowed for timed administration and removal of the solution from the wound bed. The Dakin's solution utilizing the Instill Wound V.A.C.<sup>TM</sup> was subsequently placed on his right lower extremity and his open abdomen following positive fungal cultures and concern for continued necrosis at these sites. On post injury day 16 it was also noted that he had grossly necrotic and caseating tissue in his abdomen, which was later shown to be culture positive for *Aspergillus*. The patient continued to undergo serial irrigation and debridement every 48 hours. The patient's wounds were ultimately closed without further complications on post-injury day 37.

### DISCUSSION

*Zygomycetes* and *Aspergillus* are found ubiquitously on decomposing organic material and are regularly inhaled in spore form without concern to immunocompetent patients,<sup>5,7</sup> however, IFIs continue to be among the most aggressive and devastating complications to affect our combat-injured patients. To date, we have had 77 patients diagnosed with IFI from Iraq and Afghanistan since 2001. Of these 77 patients, there have been only 4 fatalities, which is a 5.2% mortality rate compared to a reported 30% mortality rate associated with nosocomial acquired cutaneous mucormycosis following traumatic injury in the civilian literature<sup>11</sup> and the 25% mortality rate seen following the tornados in Joplin, Missouri.<sup>12</sup> Previous reports describe a mortality rate approaching 100% with disseminated angioinvasive fungal infections, such as in our case where multiple extremities and the abdomen are involved.<sup>2</sup> This high mortality rate in the civilian literature is despite utilization of aggressive debridement and appropriate antifungal therapy similar to what has been utilized in the military treatment facilities. Current systemic antifungal agents for IFI have been defined by the infectious disease literature as intravenous amphotericin B because of its in vitro activity and lower toxicity or the newer azole preparations, such as voriconazole and posaconazole in resistant cases.<sup>2,9,13,14</sup>

These infections lead to anecdotal reports of dramatic shortening, and frequently the ablation of residual limbs secondary to the

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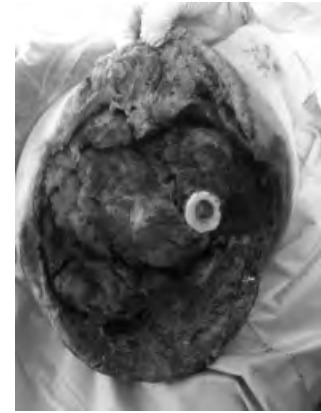
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**Figure 1.** Right elbow disarticulation following revision amputation.



**Figure 2.** Left forearm with prominent soft tissue loss and tissue necrosis.



**Figure 3.** Left transfemoral amputation with evidence of fat necrosis on presentation.

aggressive debridement of tissue necessary to gain control of the advancing necrosis. With the addition of Dakin's solution through either Instill Wound V.A.C.<sup>TM</sup> or intravenous tubing in conjunction with traditional negative pressure wound therapy, we successfully managed critically ill patients whose outcome may have been dramatically worse without this novel treatment modality.

Recently, Barsoumian, et al. have looked at the in vitro activity and toxicity of dilute Dakin's solution in comparison to mafenide acetate and amphotericin B and found that Dakin's solution possessed a more reliable efficacy and toxicity balance.<sup>15</sup> Our continued experience with Dakin's solution and initial research from the Trauma Infectious Disease Outcome Study has led to the establishment of a Joint Theater Trauma System Clinical Practice Guideline for the treatment of suspected invasive fungal infection in war wounds.<sup>16</sup> The clinical practice guidelines currently recommend commencement of Dakin's irrigation during the first debridement in theater if the patient has three of the four following risk factors, 1) dismounted blast injury, 2) above-knee immediate traumatic amputation or progressive transition from below-knee to through-knee to above-knee amputation, 3) extensive perineal, genitourital, and/or rectal injury, or 4) super massive transfusion > 25 units packed red blood cells and whole blood.<sup>17</sup>

Further direction of investigation will allow for quantitative evaluation of the loss of residual limb length and functional levels in comparison to a control group with similar blast injuries. We as well hope to define further risk factors and methods for diagnosis in order to decrease the rate of disseminated infections, and therefore decrease the morbidity and mortality of IFI.

## CONCLUSION

We have described our relative success in the treatment of IFI with the addition of Dakin's solution administration with negative pressure wound therapy as an adjunct to standard surgical management. Although this may demonstrate returning to a 150-year-old technique of wound care,<sup>10</sup> we feel that this treatment modality, in addition to aggressive debridement, and culture-directed intravenous antifungal chemotherapy can help dramatically decrease the mortality rate associated with these catastrophic infections.

This article contains the authors' most current findings, however, for a complete report of cases, refer to Lewandowski L, Purcell R, Fleming M, Gordon WT. The Use of Dilute Dakin's Solution for the Treatment of Angioinvasive Fungal Infection in the Combat Wounded: a Case Series. *Mil Med.* 2013 Apr;178(4):e503-e507.

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## Neurovascular Entrapment due to Combat-Related Heterotopic Ossification

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### INTRODUCTION

**H**eterotopic ossification (HO) is the ectopic formation of mature lamellar bone in nonosseous tissue. HO frequently develops in the upper and lower extremities following blunt trauma and is most common after elbow and acetabular fractures, burns, traumatic brain injury (TBI), and spinal cord injury.<sup>1</sup> The prevalence of HO following combat injuries, including traumatic amputations, is much higher than civilian data would suggest. Between 63% and 65% of all combat casualties with open extremity injuries or amputations develop HO.<sup>2,3</sup> In amputees, nearly 20% of affected amputations require surgical excision.<sup>3</sup> In select cases, the aberrant bone formation can envelop major neurovascular structures, such as the sciatic or tibial nerves, as well as the femoral or posterior tibial arteries, leading to symptomatic neurovascular entrapment.<sup>4</sup>

Sciatic nerve entrapment secondary to HO was first described by Kleiman and Pankovich<sup>5</sup> in 1971. Their initial case report described sciatic nerve dysfunction secondary to HO following operative fixation of a posterior fracture-dislocation of the hip. A review of the literature, summarized in Table 1, demonstrates that sciatic nerve dysfunction associated with HO is rare and most commonly reported after acetabular fractures;<sup>5-9</sup> however, two cases involved neurogenic HO<sup>10,11</sup> and another developed following repetitive trauma attributed to weight lifting.<sup>12</sup> One case report described HO leading to dysfunction of the common peroneal nerve following burn injury.<sup>13</sup> In the upper extremities, nerve entrapment by HO has been reported following trauma,<sup>14</sup> traumatic brain injury<sup>15-17</sup> and severe burns.<sup>17-19</sup> Though the relationship between systemic inflammation and HO is well-established, neurologic entrapment in the lower extremities secondary to blast trauma has not been reported to our knowledge, nor has the pre-operative planning and surgical technique been adequately discussed. Likewise, we are not aware of any reported cases of major vessel incarceration.

We present a series of five consecutive patients, two of which are detailed below, treated at our institution, who developed neurovascular entrapment secondary to severe HO after blast injury (Table 2). In doing so, we describe our preferred method of patient assessment, pre-operative planning, and surgical excision.

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### CASE SERIES

#### Case 1

A 26-year-old male sustained a blast injury due to an improvised explosive device (IED) while on foot patrol (dismounted IED). He presented with an injury severity score (ISS) of 22, mild TBI, large left anteromedial thigh soft tissue injury with volumetric muscle and soft tissue loss, as well as other minor musculoskeletal injuries to other extremities. For management of his left thigh injury, he underwent multiple irrigation and debridements until wound coverage with split-thickness skin graft (STSG). He presented at his five-month follow-up with significantly decreased knee range of motion (ROM) (0° to 15°) and focal wound ulceration over palpable, underlying HO. Pre-operative radiographs demonstrated severe HO about the medial distal femur, as shown in Figure 1. A CT angiogram demonstrated encasement of the superficial femoral vessels by HO, which was also tethering his quadriceps musculature. He returned to the operating room six months after injury for HO excision, vascular decompression, and lysis of knee adhesions with quadricepsplasty. Multiple branches of the superficial femoral artery and vein were ligated intraoperatively, but femoral and popliteal artery and vein patency and continuity were preserved. Postoperatively, the patient required additional procedures – a repeat lysis of adhesions at two months and irrigation and debridement shortly thereafter to treat a subsequent deep post-surgical infection. At final follow-up, six months after the index procedure, the patient regained active knee ROM from 0° to 90° and had no vascular symptoms. This patient, and all subsequent patients in this series, received secondary HO prophylaxis in the form of off-label use of celecoxib, 100 mg twice daily or 200 mg once daily, beginning immediately postoperative for four weeks. We observed no radiographically-evident HO recurrence in any patient.

#### Case 2

A 19-year-old male sustained a dismounted IED blast and presented with an ISS of 20, mild TBI, open distal comminuted right femur fracture, right leg soft tissue injury with posterior tibial artery transection requiring fasciotomies, and other musculoskeletal injuries. His femur fracture was initially stabilized using a knee spanning external fixator that was later converted to an intramedullary nail. His soft tissue injuries were closed with STSG following serial debridements. At his ten-month follow-up, he presented with progressive neurological deficits, continued pain, ulcerations due to underlying HO, and decreased knee and ankle ROM. He reported paraesthesias in the sural, saphenous, tibial, and superficial peroneal nerve distributions and decreased motor function (3/5 strength for tibialis anterior, extensor hallucis longus, flexor hallucis longus, and triceps surae). Pre-operative radiographs depicted in Figure 2 demonstrate moderate and severe HO about the right distal femur and tibia,



**Table 1. Previous Case Reports of Heterotopic Ossification Involving Neurologic Entrapment**

Authors	Number of patients	Inciting event	Entrapped structures	Symptoms	Electro-diagnostic studies	Treatment	Results of treatment
Kleiman, et al. <sup>5</sup>	1	Right posterior hip dislocation with posterior wall fracture s/p ORIF	Right sciatic nerve	Partial sensory and motor deficits in peroneal nerve distribution	N/A	Neurolysis and excision at 4.5 months after injury	Complete return of sensation at 2 months and no return of motor after 24 months
Derian, et al. <sup>6</sup>	1	Right posterior hip dislocation with acetabular fracture s/p closed reduction and traction	Right sciatic nerve	Complete right sciatic nerve deficit	N/A	Neurolysis and excision at 2 months after injury	Near complete return of function (all motor groups $\geq$ 4/5) 20 months post op
Manidakis, et al. <sup>8</sup>	1	Left posterior hip dislocation with posterior wall acetabular fracture s/p ORIF	Left sciatic nerve	Paraesthesiae in L5/S1 dermatomes, decreased motor function in TA and EHL	Conduction block consistent with a lesion of the sciatic nerve at the hip	Neurolysis and excision at 8 weeks	Full sensory (2 months) and motor (6 months)
Safaz, et al. <sup>11</sup>	1	TBI s/p MVA	Bilateral sciatic nerves	Weakness in bilateral lower extremities (2/5 right 4/5 left)	Bilateral axonal degeneration of the sciatic nerves (total on right, partial on left)	Physical therapy	
Jones, et al. <sup>12</sup>	1	Recurrent minor trauma from weight lifting	Left sciatic nerve	Paraesthesias of left L5/S1, complete paralysis of left TA, EHL, EDL, PL, PB weakness of GS, loss of Achilles reflex	Normal conduction of the left sciatic nerve to the level of the HO but not distal to it	Exploration and excision 2 months after symptoms	At 24 months, no detectable motor weakness, improved but decreased sensation, no improvement in Achilles reflex
Thakker, et al. <sup>9</sup>	1	Left posterior hip dislocation with acetabular fracture and sciatic nerve laceration s/p ORIF and nerve repair	Left sciatic nerve	Pain and paraesthesias along medial aspect of his leg and foot drop	None	Exploration, excision, neurolysis	Resolution of pain and paraesthesias with continued foot drop at 2 years post op
Brooke, et al. <sup>21</sup>	1	Closed head injury after motorcycle accident	Right femoral nerve	0/5 motor in the quadriceps; decreased hip ROM	4+ positive sharp waves and fibrillations without voluntary motor units in the quadriceps	Exploration and excision of HO with epineuriolysis	Improved hip ROM with near normal return of motor function at 19 months
Laborde, et al. <sup>10</sup>	1	Respiratory distress and intubation x 3 weeks	Right sciatic nerve	Weakness in right leg in the tibial nerve distribution	N/A	Medical management with cox-2 inhibitors	Not reported

s/p, status post; ORIF, open reduction and internal fixation; TA, tibialis anterior; EHL, extensor hallucis longus; TBI, traumatic brain injury; MVA, motor vehicle accident; EDL, extensor digitorum longus; PL, peroneus longus; PB, peroneus brevis; GS, gastroc-soleus

**Table 2. Description of Cases**

Case	Age	ISS	TBI	Injury leading to HO	Grade of HO	Symptoms	Enveloped structures	Post surgery
1	26	22	Mild	Left thigh STI	Severe	Left knee decreased ROM (flexion-extension arc 0°-15°), ulceration, pain	Superficial femoral vessels	Improved active knee ROM (flexion-extension arc 0°-140°), resolution of ulcerations, and no vascular compromise
2	19	20	Mild	Open right femur fracture and right leg STI	Moderate (femur); severe (tibia)	Foot paraesthesias and decreased motor function (3/5 TA, EHL, FHL, and GS)	Profunda femoris artery and vein, posterior tibial and peroneal vessels, tibial nerve	Improvement in sensation and pain, resolution of paraesthesias, regained full motor strength
3	22	29	Mild	Open right femur fracture	Severe	Right thigh discomfort, decreased knee ROM plantar paraesthesias	Sciatic nerve	Regained full knee ROM; paraesthesias resolved
4	25	29	Mild	Right thigh STI	Severe	Progressive neurologic deficits (3/5 TA, 2/5 GS, 2/5 HS), ulceration, pain, and decreased R hip ROM (10°-50° flexion, 10° adduction, 15° abduction)	Sciatic nerve	Improved hip ROM, distal dysesthesias, and motor function (3/5 TA, 4/5 GS, 3/5 HS) improved
5	27	29	Mild	Left transfemoral amputation	Severe	Decreased left hip ROM (4° adduction, 0°-10° hip flexion-extension arc), pain	Sciatic nerve	Improvement in hip ROM (flexion-extension arc 0-90° actively and 0-105° passively)

ISS, injury severity score; TBI, traumatic brain injury; HO, heterotopic ossification; STI, soft tissue injury; ROM, range of motion; TA, tibialis anterior; EHL, extensor hallucis longus; FHL, flexor hallucis longus; GS, gastrocnemius-soleus complex; HS, hamstrings



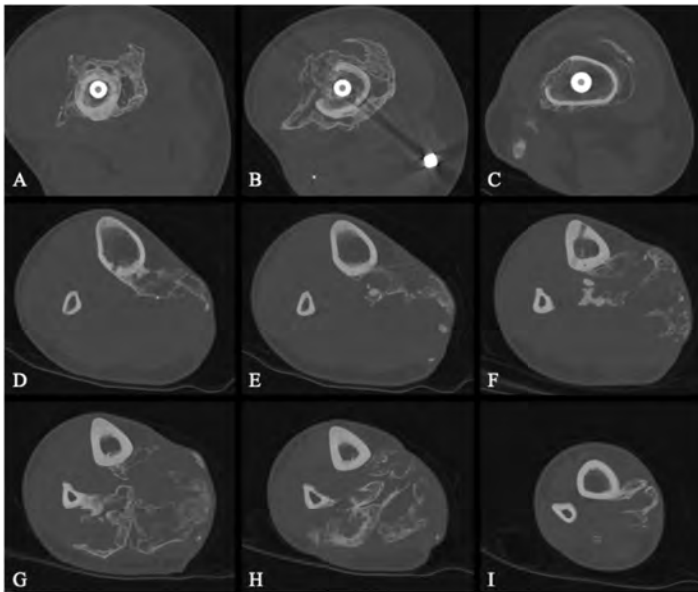
**Figure 1.** Pre-operative AP (A) and lateral (B) radiographs of the left distal femur of case 1 demonstrating severe HO.



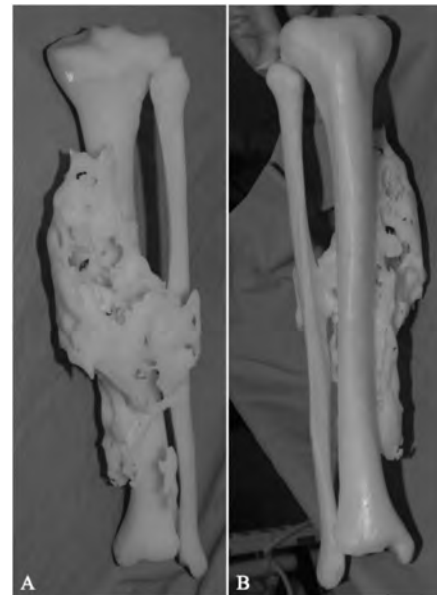
**Figure 2.** Pre-operative AP (A) and lateral (B) radiographs of the right distal femur, demonstrating moderate HO and AP (C) and lateral (D) radiographs of the right tibia, demonstrating severe HO in case 2.

respectively. A CT scan demonstrated encasement of the profunda femoris artery and vein and the posterior tibial and peroneal vessels and the tibial nerve in the leg, shown in Figure 3. The CT scan was used to make a resin model, seen in Figure 4. Fourteen months after injury, he underwent HO excision of the right posterior thigh and leg,

as well as excision of the entire STSG and overlying areas of ulceration. Intra-operative photographs are seen in Figure 5. Following excision, he reported immediate improvement in sensation and pain, and complete resolution of paraesthesias. At his 11-month follow-up, he had full motor strength in all distributions.



**Figure 3.** Axial CT scan, moving from proximal to distal, of case 2, demonstrating encasement of the deep artery and vein in the thigh (A-C) and the posterior tibial and peroneal arteries as well as the tibial nerve in the leg (D-I).



**Figure 4.** Posterior (A) and anterior (B) views of the 3D resin model constructed from the CT scan of case 2 and used intraoperatively.



**Figure 5.** Intraoperative images demonstrating the initial mass of HO with proximal and distal identification of the neurovascular bundle marked with vessel loops (A), after removal of a portion of the HO that was directly over the neurovascular bundle (B). Final image after removal of the symptomatic HO (C).

## DISCUSSION

The cases presented herein demonstrate that severe heterotopic ossification can lead to late neurologic and vascular symptoms due to neurovascular entrapment. These symptoms can be seen in addition to the more commonly described sequelae of HO, such as pain, wound ulceration, and decreased joint ROM. All five patients sustained blast injuries and mild TBI. During outpatient rehabilitation, each developed debilitating symptoms that halted, and in some cases regressed, their progress with therapy. The patients returned to the operating room, on average, ten months after initial injury for HO excision, decompression or neurolysis of the entrapped neurovascular structures, and concurrent procedures such as soft tissue revisions or lysis of adhesions.

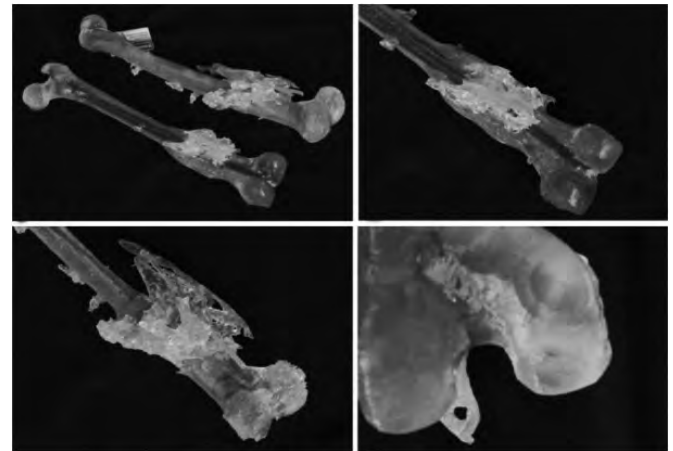
Previous reports of HO with symptomatic neurologic entrapment have focused predominantly on reporting singular cases and clinical outcomes, with respect to resolution of neurologic symptoms and recurrence, as opposed to the pre-operative planning and surgical techniques, which are the focus of this report. However, our clinical outcomes globally resulted in successful functional preservation of the entrapped neurovascular structures and improved ROM without clinical or radiologic evidence of HO recurrence.

## Preoperative Planning

Preoperative assessment begins with a thorough physical examination. Neurological deficits should be characterized as static or progressive and focus on the distribution of the sensory deficits and degree of motor weakness. The soft tissue envelope should be examined for previous traumatic wounds, surgical incisions, STSG, and ulcerations that require excision. We obtain pre-operative CT scans to characterize the HO and the relationship between the HO and named neurovascular structures, as recommended.<sup>4,7,8,20</sup> CT angiograms can also yield important information when there is concern for vessel involvement.

At our institution, CT also allows for the construction of three-dimensional resin models, as seen in Figure 6. These models allow one to plan the operative approach and osteotomies, as well as to refer to the model intraoperatively. We find this helpful because each lesion is distinct and uniquely shaped, and the lesional topography becomes particularly important given the obfuscation of normal anatomy common in blast wounds that have undergone various types of surgical procedures, as well as tethering of structures to and within the HO. Intraoperatively, these models are referenced as anatomical guides to dissection and piecemeal resection.

Each patient is typed and cross-matched for at least two units of packed red blood cells. Additionally, we find the following equipment helpful during surgery: loupe magnification, bipolar coagulation in addition to standard monopolar electrocautery, a full array of ronguers and periosteal elevators (*e.g.*, Cobb, Key), as well as instruments and available expertise necessary to reconstruct/repair the involved nerve(s) or vessel(s)—although the latter have fortunately not been necessary to date. Koulouvaris and colleagues<sup>20</sup> also include cell saver and possible intra-operative electrical stimulation, which we have utilized twice, in their preparations for surgery as well as a pre-operative bone scan to determine the maturity of the HO prior to excision. There is no clear consensus on when ectopic bone lesions should be excised, however, in the setting of progressive neurological deficits and symptoms, early surgical exploration/excision of HO is advocated.<sup>5,6,8,21</sup>



**Figure 6.** Examples of two resin models demonstrating heterotopic ossification of the femur. They were used to plan the operative approach and as anatomical guides to dissection and piecemeal resection.

## Operative Technique

Excision of heterotopic ossification is associated with relatively high rates of major short-term complications, including neurovascular injury, substantial blood loss, increased hospital stay, fracture, and infection.<sup>20,22</sup> We attempt to use prior traumatic incisions whenever possible. If this is not a viable option, we use a direct approach over the most superficial symptomatic area and extend our exposure longitudinally from that region. The involved neurovascular structures are identified and isolated both proximal and distal to the area of entrapment whenever remotely feasible. Once adequate exposure or limited removal of the HO mass has been achieved, the nerve or vessels can be traced directly into the offending area and gradually mobilized via meticulous sharp and blunt dissection and pituitary rongeur removal of the overlying HO. Kleiman and coauthors<sup>5</sup> described removal of the entire mass of HO, but partial resection of only the symptomatic areas can also be successful in carefully selected patients.<sup>3,4,20</sup> Our aim is not necessarily to remove all HO if doing so would sacrifice a substantial amount of uninvolved soft tissue, although complete resection is performed when practicable. In these latter instances, we remove only that which is symptomatic—in this series, the HO that was entrapping/compressing the neurovascular structures.

Post-operative secondary prophylaxis has also been described such as the use of nonsteroidal anti-inflammatory drugs (indomethacin,<sup>20,23,24</sup> cyclooxygenase-2 inhibitors, naproxen, ibuprofen), local radiation therapy<sup>25,26</sup> (700 cGy preoperatively or within 24 hours post-operatively), a combination of both,<sup>3,4,8,27,28</sup> and no prophylaxis. In the present series, each patient received celecoxib, as described above, and no patient developed radiographically evident or symptomatic recurrence of HO.

In the short term, all patients who underwent excision demonstrated functional improvement. All patients who had complaints of neuropathic pain had improvement in their pain after the initial post-operative phase. Those with impaired ROM regained motion as described above. As might be expected, we found that sensory deficits resolved prior to motor deficits.

Long-term complications of HO excisions include recurrence and wound breakdown. Heterotopic ossification recurrence has been described in the literature with varying rates, from zero to nearly 100%;<sup>22-25</sup> however, the severity of recurrence was not always described. Furthermore, the recurrence rate varies widely with the inciting event (e.g., higher recurrence rates are typically associated with neurogenic HO), timing of excision, and secondary prophylaxis. Ebinger, et al.<sup>26</sup> reported a recurrence rate of 93% in the hip, although none of their patients required repeat excision, whereas Genet, et al.<sup>28</sup> had a recurrence rate of 6%, but only reported patients requiring repeat excision. A meta-analysis<sup>27</sup> evaluating recurrence of HO after TBI included 255 patients in 16 studies. The authors estimated the recurrence rate at 19.8%, but did not describe the clinical relevance of these recurrences. Fortunately, clinically significant HO recurrence has not been a frequent complication in combat casualties<sup>3,29</sup> and did not occur in this series. In the present series, however, one patient developed wound-healing complications leading to infection and operative intervention.

In conclusion, we found that HO entrapping major neurovascular structures following severe extremity trauma can be safely excised with careful pre-operative planning and meticulous operative excision. Excision is associated with a relatively low rate of recurrence and reliably appears to resolve the symptoms associated with neurovascular compression. Generally favorable clinical results are achievable despite the relative complexity of the dissections and excisions required.

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## Outcomes of Single-Level Cervical Disc Arthroplasty versus Anterior Cervical Discectomy and Fusion: a Single Center, Retrospective Review

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### INTRODUCTION

Cervical disc arthroplasty (CDA) has been espoused as a safe, segmental motion-sparing alternative to anterior discectomy and fusion (ACDF) in the treatment of cervical radiculopathy and myelopathy resulting from spondylosis and acute disc herniation. Although ACDF is the current standard for treatment of cervical radiculopathy and myelopathy, concerns exist over symptomatic adjacent-segment disc degeneration.<sup>1-2</sup> Furthermore, cadaver studies have demonstrated increased motion and intra-discal pressures adjacent to cervical fusion levels.<sup>3-4</sup> These concerns have led to the development and use of several cervical disc arthroplasty systems. Anticipated benefits of CDA include maintaining alignment and motion as well as decreasing stress on the adjacent-level disc, which may decrease symptomatic adjacent level disc degeneration.<sup>5</sup> General indications for CDA include reconstruction after neural decompression of disc herniation or foraminal osteophytes causing radiculopathy or myelopathy. Contraindications include deformity, immobile segments, instability, and facet joint degeneration. Relative contraindications include rheumatoid arthritis, renal failure, osteoporosis, cancer, and pre-operative corticosteroids.<sup>6-7</sup>

Short-term results from several small randomized controlled trials demonstrated that CDA was associated with better overall success, better neurologic success, and fewer revision procedures.<sup>8-12</sup> However, there are few single center comparison studies, and current studies do not contain large numbers of patients. Based on our experience with CDA, we performed a review comparing outcomes and complications in patients undergoing single-level CDA and single-level ACDF at a single institution.

### MATERIALS AND METHODS

Following approval from our institutional review board, the surgical database at our institution was queried to identify all patients who had undergone single-level cervical disc arthroplasty or single-level anterior discectomy and fusion between August 2008 and August 2012. Seven surgeons (5 neurological surgeons and 2 spine fellowship-trained orthopaedic surgeons) were the primary surgeon performing the respective procedure. The search yielded 180 CDA patients and 95 ACDF patients. In the CDA group, 9 patients were lost to follow-up and

7 patients in the ACDF group were also lost to follow-up, which left 259 total patients (171 in the CDA group and 88 in the ACDF group) for review. Both groups included patients undergoing primary and revision surgery. All data were collected via a retrospective chart analysis, which included inpatient and outpatient clinical notes, surgical databases, and radiographs, and subsequently analyzed by independent researchers. Data collected included patient demographic information (age, sex, tobacco use, BMI), patient-centered outcomes (complete relief of pre-operative symptoms, relief of pre-operative neurologic symptoms, return to pre-operative level of activity, return to active duty for patients in the military), and complications (incidence of persistent post-operative posterior neck pain, recurrent laryngeal nerve injury, persistent dysphagia, post-operative respiratory compromise, esophageal/tracheal disruption, implant failure, adjacent segment degeneration, intra-operative fracture, dural tear, nerve root injury). Persistent posterior neck pain and dysphagia were defined in the study as symptoms lasting longer than 3 months in the post-operative period or requiring secondary intervention.

### RESULTS

In the CDA group, there were 134 males (78.4%) and 37 females (21.6%) with an average age of 40.5 ( $\pm$  8.3) years. In the ACDF group, there were 65 males (73.9%) and 23 females (26.1%) and the average age was 47.5 ( $\pm$  12.4) years. The average follow-up was 9.8 ( $\pm$  9.9) months for the CDA group and 11.8 ( $\pm$  9.6) months for the ACDF group. The average BMI was 27.8 ( $\pm$  3.8) kg/m<sup>2</sup> for CDA patients and 28.9 ( $\pm$  5.7) kg/m<sup>2</sup> for ACDF patients. Tobacco use was higher in the CDA group at 28.1% (132 patients) as compared to the ACDF group at 17.0% (15 patients). Revision surgery was more common in the ACDF group than the CDA group (23.9% versus 4.7%). In the CDA group, 132 patients (77.2%) were on active duty in the military at the time of surgery compared to only 40 patients (45.5%) in the ACDF group.

The primary indication for CDA was radiculopathy in 154 patients (90.1%). Other indications included myelopathy (1.8%), myelo-radiculopathy (5.8%), and neck pain (2.3%). In the ACDF group, the primary indication for surgery was radiculopathy in only 63.6% of patients and myelopathy in 18.2% of patients. Other indications included myelo-radiculopathy (5.7%), neck pain (5.7%), trauma (3.4%), and pseudoarthrosis (3.4%). The most common levels addressed at the time of surgery in the CDA group were C6-7 followed by C5-6 (50.9% and 38.0%, respectively). In the ACDF cohort, the most common levels of disease were C5-6 and C6-7 (44.3% and 23.9%, respectively). Levels C3-4 and C4-5 were commonly involved in the ACDF group (13.6% and 15.9%, respectively) as compared to the CDA group (2.3% and 8.2%, respectively for C3-4 and C4-5). One patient underwent anterior cervical discectomy and fusion at C2-3 and one patient in each group had surgery at C7-T1 (Table 1).

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**Table 1. Patient Demographics**

	CDA	ACDF
Total patients	171	88
Males	134 (78.4%)	65 (73.9%)
Females	37 (21.6%)	23 (26.1%)
Age	40.5 ± 8.3 years	47.5 ± 12.4 years
BMI	27.8 (± 3.8) kg/m <sup>2</sup>	28.9 (± 5.7) kg/m <sup>2</sup>
Tobacco use	48 (28.1%)	15 (17.0%)
Active duty military	132 (77.2%)	40 (45.5%)
Revision surgery	8 (4.7%)	21 (23.9%)
Average follow-up	9.8 ± 9.9 months	11.8 ± 9.6 months
Levels of disease		
C2-3	0	1 (1.1%)
C3-4	4 (2.3%)	12 (13.6%)
C4-5	14 (8.2%)	14 (15.9%)
C5-6	65 (38.0%)	39 (44.3%)
C6-7	87 (50.9%)	21 (23.9%)
C7-T1	1 (0.6%)	1 (1.1%)
Primary indication for surgery		
Myelopathy	3 (1.8%)	16 (18.2%)
Radiculopathy	154 (90.1%)	56 (63.6%)
Myelo-radiculopathy	10 (5.8%)	5 (5.7%)
Neck pain	4 (2.3%)	5 (5.7%)
Trauma	N/A	3 (3.4%)
Pseudoarthrosis	N/A	3 (3.4%)

CDA, cervical disc arthroplasty; ACDF, anterior discectomy and fusion; N/A, not applicable

The Prestige™ (Medtronic, Memphis, TN) cervical arthroplasty system was utilized in the majority of patients (94.7%), while the ProDisc-C system (Depuy Synthes, Paoli, PA) was utilized in the remainder of patients (5.3%). The most common surgical implant used in the ACDF group was the traditional anterior cervical plate and interbody spacer (78.4%). Other implants included the stand alone integrated spacer in 11.4% of patients and a spacer-only construct in 10.2% of ACDF patients (Table 2).

**Table 2. Type of Surgical Implant**

Surgical Implant	CDA	ACDF
Prestige	162 (94.7%)	N/A
Pro-Disc C	9 (5.3%)	N/A
Anterior plate/spacer	N/A	69 (78.4%)
Stand alone integrated spacer	N/A	10 (11.4%)
Spacer only	N/A	9 (10.2%)

CDA, cervical disc arthroplasty; ACDF, anterior discectomy and fusion; N/A, not applicable

Of the 171 patients in the CDA group, 157 (91.8%) experienced complete relief of pre-operative neurologic symptoms and 154 patients (90.1%) had complete post-operative symptomatic relief. In the ACDF group, 78 (88.6%) patients had post-operative relief of neurologic symptoms and 76 (86.4%) patients had complete relief of all pre-operative symptoms. Seventeen patients (9.9%) in the CDA group and 12 patients in the ACDF group (13.6%) had incomplete relief of pre-operative symptoms. These patients had persistent posterior neck pain, continued radiculopathy, continued myelopathy, or a combination of the aforementioned symptoms. In both groups, a vast majority of patients were able to return to their pre-operative level of activity (93.0% in the CDA group and 88.6% in the ACDF group). In the active duty military population, 92.4% of patients in the CDA and 84.0% of patients in the ACDF group were able to return to active duty after surgery (Table 3).

Persistent posterior neck pain was observed post-operatively in both cohorts at rates of 15.4% (27 patients) in the CDA group and 12.5% (11 patients) in the ACDF group. Twenty patients (74.1%) in the CDA group who experienced persistent posterior neck pain post-operatively eventually had resolution of their symptoms without intervention at the last recorded follow-up appointment. Similarly, 5 patients (45.5%) in the ACDF group also had resolution by their last recorded follow-up visit. The complication rate was 9.9% in the CDA group, which included recurrent laryngeal nerve injury (2.9%), persistent dysphagia (5.8%), superficial infection (0.6%), and nerve root injury (0.6%). The ACDF complication rate was 9.1%, which included persistent dysphagia (3.4%), pseudoarthrosis (3.4%), spinal cord injury (1.1%), and dural tear (1.1%). No implant failures, episodes of post-operative respiratory compromise, esophageal/tracheal disruptions, nor intra-operative fractures were observed in either group in our study. The reoperation rate in the CDA group was 3.5%, including posterior decompression and fusion, adjacent segment degeneration requiring surgery, and conversion of CDA to ACDF. In the ACDF group, the reoperation rate was 5.7%, including posterior decompression and fusion and revision ACDF (Table 4).

## DISCUSSION

This study represents the largest, non-funded retrospective comparison review of single-level CDA and single-level ACDF. We found a 90.1% rate of complete pre-operative relief of symptoms and

**Table 3. Surgical Outcomes**

Outcomes	CDA	ACDF
Relief of neurologic symptoms	157 (91.8%)	78 (88.6%)
Complete relief of symptoms	154 (90.1%)	76 (86.4%)
Return to full activity	159 (93.0%)	78 (88.6%)
Return to active duty military	122/132 (92.4%)	32/40 (84.0%)

CDA, cervical disc arthroplasty; ACDF, anterior discectomy and fusion

**Table 4. Surgical Complications**

Complications	CDA	ACDF
Persistent posterior neck pain	27 (15.4%)	11 (12.5%)
Recurrent laryngeal nerve injury	5 (2.9%)	0 (0.0%)
Persistent dysphagia	10 (5.8%)	3 (3.4%)
Infection (superficial)	1 (0.6%)	0
Pseudoarthrosis	N/A	3 (3.4%)
Nerve root injury	1 (0.6%)	0
Spinal cord injury	0	1 (1.1%)
Dural tear	0	1 (1.1%)
Reoperation during follow-up	6 (3.5%)	5 (5.7%)
Posterior decompression/fusion	2 (1.2%)	3 (3.4%)
Adjacent segment degeneration	2 (1.2%)	0
Conversion to ACDF	2 (1.2%)	N/A
Revision ACDF	N/A	2 (2.3%)

CDA, cervical disc arthroplasty; ACDF, anterior discectomy and fusion; N/A, not applicable

93.0% rate of return to pre-operative level of activity in the CDA group, which compared to a complete pre-operative symptomatic relief rate of 86.4% and a return to full activity rate of 88.6% in the ACDF group. Active duty service members included in the study returned to active duty post-operatively at a rate of 92.4% in the CDA group and 84.0% in the ACDF group. The disparity in complete symptomatic relief and return to full activity between the groups can partially be attributed to differences in patient demographics. The percentage of patients undergoing revision was five times higher in the ACDF group. Additionally, the indication for surgery for over 90% of patients in the CDA group

was radiculopathy, compared to only 64% of patients in the ACDF group.

This review also demonstrated a low complication rate for both groups, in particular for recurrent laryngeal nerve injury (2.9% in the CDA group, 0% in the ACDF group), persistent dysphagia (5.8% in the CDA group, 3.4% in the ACDF group), and adjacent segment degeneration requiring surgery (1.2% in the CDA group, 0% in the ACDF group). The dysphagia rate observed for both groups was lower than traditionally reported in the literature for either single-level CDA or single-level ACDF.<sup>13-17</sup> Reoperation rates were lower in the CDA group compared to the ACDF group. These results are consistent with the rates observed in recent randomized controlled trials between CDA and ACDF.<sup>9, 11</sup>

While short-term results have demonstrated CDA as a safe and reliable alternative to ACDF, no long-term results currently exist comparing these two surgical techniques. At this point, ACDF continues to be the most common surgical treatment for cervical radiculopathy and other degenerative conditions in the neck.<sup>18-19</sup> Before CDA can be routinely implemented into clinical practice, studies must prove it to be at least as effective in symptomatic relief as ACDF, with similar or lower complication rates, and to reduce the incidence of adjacent segment degeneration. However, two recent prospective, multi-center, randomized, controlled Food and Drug Administration (FDA) Investigational Device Exemption (IDE) studies between single-level CDA and single-level ACDF with 48-month and 60-month follow-up, respectively, show promising mid-term results for two different cervical disc arthroplasty systems. The CDA groups demonstrated a higher rate of overall success, greater improvements in neck disability index (NDI), neck and arm pain scores, and SF-36 Physical Component Scores as compared with patients in the ACDF group. The adjacent segment degeneration rate was lower at 60 months for the CDA group. Additionally, normal segmental motion was maintained in the CDA group. Finally, revision rates and supplemental fixation surgical procedures were lower following CDA.<sup>20-21</sup> These two studies again demonstrate that CDA is a viable treatment option for cervical radiculopathy and myelopathy.

To our knowledge, this review represents the largest, non-funded, comparison study between single-level CDA and single-level ACDF. In the CDA group, we found a 90% rate of complete pre-operative symptom relief, with 93% of patients returning to their pre-operative level of activity, compared to rates 86% and 89%, respectively, in the ACDF group. Complication rates in both groups were lower than reported in the literature. This study demonstrates that CDA is a safe and reliable alternative to ACDF, but long-term studies are needed to ascertain differences in outcomes, complication rates, and adjacent segment degeneration.

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**Cervical Posterior Foraminotomy's Effect on Segmental Range of Motion in the Setting of Total Disc Arthroplasty**

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**INTRODUCTION**

Cervical disc arthroplasty (CDA) is being used with increasing frequency for the treatment of cervical radiculopathy.<sup>1,2</sup> Prospective randomized data has emerged over the past few years indicating that outcomes of CDA are comparable to anterior cervical discectomy and fusion (ACDF), which has long been considered the historic gold standard.<sup>3-7</sup> As CDA becomes more common and the number of patients living with a cervical disc implant increases, so will complications and the need to perform revision surgery.<sup>8,9</sup> Recurrent radiculopathy is a leading cause of revision surgery for both ACDF and CDA. In the setting of ACDF, recurrent symptoms can be secondary to inadequate nerve root decompression, pseudoarthrosis, or implant failure. The surgical revision strategy may involve a repeat anterior approach, or decompression can be performed via a posterior laminoforaminotomy. In itself, posterior laminoforaminotomy has a proven track record and is an effective alternative to anterior procedures for primary cervical radiculopathy.<sup>10-13</sup> When posterior foraminotomy is executed in conjunction with a stable ACDF, the combination of procedures carry a minimal risk of iatrogenically induced instability. Conversely, ACDF can be safely performed as a revision procedure for recurrent symptoms following a primary posterior foraminotomy.

In comparison, CDA differs from ACDF in that it is not a fusion procedure and its intent is to maintain segmental cervical spine motion.<sup>14,15</sup> Furthermore, posterior foraminotomy can increase cervical spine range of motion (ROM), and potentially induce cervical spine instability.<sup>13</sup> The question that is unknown is whether CDA and foraminotomy can be safely performed without causing excessive cervical spine motion or iatrogenic instability. This point serves as the impetus for the current study, as we sought to investigate the effect on segmental cervical spine motion of CDA performed in combination with posterior foraminotomy.

**METHODS**

Following institutional review board (IRB) approval, 10 fresh human cadaveric spine specimens from occiput to sacrum were obtained

from the Maryland State Anatomy Board. The medical history for each cadaver was reviewed and specimens were excluded for a history of primary or secondary bone disease. Upon receipt of the specimens, all muscular attachments were dissected free from the spinal column, with care taken to preserve the osseous and discoligamentous structures. Specimens were stored at -20° C until biomechanical testing was initiated, at which time they were thawed to room temperature. Prior to testing, dual X-ray absorptiometry (DEXA) using Hologic QDR-2000 (Boston, MA) was performed on each specimen to document pre-testing bone mineral density. Additionally, plain X-rays were taken to ensure normal gross anatomy and the absence of destructive bone lesions.

Each specimen was sectioned at the occ-C1 level cephalad and the T3-T4 level caudally. They were then secured using a K-wire fixation jig and polyester resin, proximally at C1 and distally at T3. Biomechanical testing was performed utilizing a MTS 858 Bionix Testing System configured with a custom-built 6 Degree-of-freedom Spine Simulator (MTS Corporation, Minneapolis, MN), allowing pure, unconstrained multi-directional load application. Intersegmental motion evaluation was analyzed with the use of specialized markers consisting of three non-co-linear infrared light emitting diodes (LEDs). One marker was rigidly attached to each vertebral level (C3-C7) and oriented to permit detection by an optoelectronic motion analysis system (OptoTrak Certus, Northern Digital Inc., Waterloo, Ontario, Canada). The specimens were non-destructively tested under axial rotation (Y-axis,  $\pm 2$  Nm), flexion/extension (X-axis,  $\pm 2$  Nm), and lateral bending (Z-axis,  $\pm 2$  Nm) testing modes using a pure moment loading system. Each test was repeated for two loading and unloading cycles, with data from the second cycle used for computational analysis. For non-destructive multi-directional flexibility analysis, the peak range of motion for each loading mode was calculated as the sum of motions [maximum  $\pm$  rotation for torsion, flexion-extension and left + right bending (degrees)] observed in the neutral (NZ) and elastic (EZ) zones at the second loading cycle (ROM = NZ + EZ).

The specimens were initially tested, according to the above protocol, in the intact state (group 1) to establish baseline (control) ROM values. Each specimen was then sequentially tested four more times according to the following experimental groups: 2) C5-6 CDA, 3) C5-6 CDA with unilateral C5-6 foraminotomy, 4) C5-6 CDA with bilateral C5-6 foraminotomy, 5) C5-6 CDA with bilateral C5-6 foraminotomy and bilateral C4-5 foraminotomy.

The Prestige Cervical Disc (Medtronic, Memphis TN) served as the arthroplasty device for the study. The implants were placed following a complete C5-C6 discectomy in accordance with the manufacturer technique guide. The posterior foraminotomies were performed first by identifying the location of the cervical facet joint in the poste-

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rior cervical spine. Approximately one-third of the cephalad lamina just medial to the inferior facet was resected with a high speed bur. The lateral aspect of the ligamentum flavum and remaining soft tissues were then resected with a Kerrison rongeur. From this point, the inferior medial half of the inferior articular facet was thinned and resected to expose the underlying superior articular facet. Once the medial half of the superior facet was sufficiently visualized, the superior and medial half of the facet was thinned and resected using a high speed bur and Kerrison rongeur. Care was taken to avoid resecting greater than 50% of the facet joint. A small nerve hook was utilized to ensure adequate decompression through direct palpation of the superior and lateral pedicle walls. When the lateral pedicle wall was easily reached by the probe, the posterior foraminotomy was considered complete.

ROM data was reported as degrees. The mean values of the intact group and each experimental group was defined in terms of axial rotation (AR), lateral bending (LB), and flexion/extension (FE) for each cervical segment from C3-C7. Statistical analysis was performed using SPSS version 18.0 software (SPSS, Inc., Chicago, IL) and included a repeated-measures analysis of variance (ANOVA), followed by a test of simple effects as a post-hoc intergroup evaluation using Sidak correction for multiple comparisons. Statistical results at  $p < 0.05$  were considered significant.

## RESULTS

A total of seven specimens underwent complete ROM testing and were included in the final statistical analysis. Of the ten specimens initially selected for study, one was eliminated secondary to iatrogenic fracture caused during fixation to the biomechanical testing jig, and two others were eliminated for ROM values that were greater than two standard deviations outside the mean.

ROM of the C5-C6 spinal segment was most affected across the experimental groups and within this segment; the flexion/extension plane experienced the greatest increases. Flexion and extension increased from 10.7° in the intact specimens (group 1) to 14.1° in the CDA specimens (group 2) and further to 16° to 17° in the CDA + foraminotomy groups (groups 3 to 5). The ROM difference between group 1 (intact/control) and groups 3 to 5 (CDA + foraminotomy) was statistically significant ( $p < 0.007$ ). However, there was not a significant ROM increase between group 1 (intact) and group 2 (CDA),  $p > 0.2$ . Furthermore, there was no significant difference in ROM between group 2 (CDA) and groups 3 to 5 (CDA + foraminotomy),  $p > 0.3$ . A stepwise increase in axial rotation was observed from group 1 (7.6°) to group 5 (10.9°), although this increase was not statistically significant,  $p > 0.3$ . The lateral bending plane experienced the smallest ROM changes remaining relatively stable across the tested groups, yielding 7.5° in group 1 and increasing to 7.9° in group 5,  $p > 0.9$  (see Table 1).

There were relatively small and universally insignificant ROM changes at the C4-5 and C6-7 spinal segments. At C4-5, flexion/extension had the largest change in ROM, increasing from 9.9° in group 1 (intact) to 13.2° in group 5 (CDA + C56 and C45 foraminotomy), but the difference was not statistically significant,  $p > 0.8$ . Axial rotation and lateral bending had minimal changes at C4-5 segment that did not reach statistical significance,  $p > 0.6$ . The C6-7 segment had a small but insignificant decrease in axial rotation from group 1 to groups 2 to 5,  $p > 0.8$ . Both lateral bending and flexion/extension remained relatively stable across all groups (1 to 5) at C6-7,  $p > 0.7$  (see Tables 2 and 3).

Table 1. C5-C6 Range of Motion				
Group		Axial rotation	Lateral bending	Flexion extension
1	Intact	7.6 ± 2	7.5 ± 3	10.7 ± 3
2	C5-6 CDA	8.8 ± 2	6.6 ± 2	14.1 ± 4 ▲
3	C5-6 CDA with unilateral C5-6 foraminotomy	9.8 ± 4	7.4 ± 3	16.7 ± 2* ▲
4	C5-6 CDA with bilateral C5-6 foraminotomy	9.5 ± 4	7.5 ± 3	17.2 ± 2* ▲
5	C5-6 CDA with bilateral C5-6 foraminotomy + bilateral C4-5 foraminotomy	10.9 ± 4	7.9 ± 3	16.7 ± 3* ▲
	$\alpha$	$p > 0.3$	$p > 0.9$	$p < 0.007$
*, significantly different compared to intact (control) state; ▲, no significant difference between experimental groups				

Table 2. C4-C5 Range of Motion				
Group		Axial rotation	Lateral bending	Flexion extension
1	Intact	8.8 ± 5	8.5 ± 3	9.9 ± 3
2	C5-6 CDA	9.6 ± 5	8.3 ± 4	13.4 ± 7
3	C5-6 CDA with unilateral C5-6 foraminotomy	8.1 ± 3	9.4 ± 4	12.4 ± 7
4	C5-6 CDA with bilateral C5-6 foraminotomy	8.7 ± 4	9.7 ± 4	12.8 ± 6
5	C5-6 CDA with bilateral C5-6 foraminotomy + bilateral C4-5 foraminotomy	8.8 ± 4	10.8 ± 4	13.2 ± 7
	$\alpha$	$p > 0.9$	$p > 0.6$	$p > 0.8$

Combined range of motion across all measured levels, reported as C4-7, did not significantly change across the tested groups. Both axial rotation and lateral bending ROM values remained relatively stable. Axial rotation increased from 23.5° in group 1 to 25.9° in group 2 ( $p > 0.9$ ) and lateral bending increased from 24.3° in group 1 to 26.8° in group 5 ( $p > 0.9$ ). In the flexion/extension plane there was a 7.5° increase from group 1 to 2 (31.4°→38.9°) and although the change was not significant,  $p > 0.1$ , it indicates that placement of the CDA at C56 had the greatest impact on motion in this plane.

Table 3. C6-C7 Range of Motion				
Group		Axial rotation	Lateral bending	Flexion extension
1	Intact	7.1 ± 3	8.4 ± 4	10.7 ± 2
2	C5-6 CDA	5.9 ± 4	8.4 ± 4	11.3 ± 2
3	C5-6 CDA with unilateral C5-6 foraminotomy	5.3 ± 2	8.2 ± 4	11.2 ± 2
4	C5-6 CDA with bilateral C5-6 foraminotomy	5.8 ± 3	8.1 ± 4	11.9 ± 2
5	C5-6 CDA with bilateral C5-6 foraminotomy + bilateral C4-5 foraminotomy	6.2 ± 2	8.1 ± 3	12.1 ± 2
	$\alpha$	$p > 0.8$	$p > 0.9$	$p > 0.7$

Table 4. C4-C7 Range of Motion				
Group		Axial rotation	Lateral bending	Flexion extension
1	Intact	23.5 ± 9	24.3 ± 9	31.4 ± 5
2	C5-6 CDA	22.9 ± 9	23.3 ± 7	38.9 ± 10
3	C5-6 CDA with unilateral C5-6 foraminotomy	23.1 ± 9	25.1 ± 9	40.4 ± 9
4	C5-6 CDA with bilateral C5-6 foraminotomy	24.0 ± 10	25.3 ± 9	41.8 ± 9
5	C5-6 CDA with bilateral C5-6 foraminotomy + bilateral C4-5 foraminotomy	25.9 ± 7	26.8 ± 9	41.9 ± 9
	$\alpha$	$p > 0.9$	$p > 0.9$	$p > 0.1$

Flexion extension remained stable with minimal increase from group 2 to group 5 (38.5°→41.9°) (see Table 4).

## DISCUSSION

The addition of a posterior foraminotomy to cervical spine specimens implanted with a CDA did not cause a statistically significant increase in segmental ROM. This finding held true in the setting of bilateral foraminotomies, and even when foraminotomies were added to a supra-adjacent level. The flexion/extension plane experienced the largest ROM changes and a significant increase was observed when

comparing the intact specimens to those that had both foraminotomy and CDA. Lateral bending and axial rotation were minimally affected by either CDA or the addition of foraminotomies. Flexion and extension at the C56 level increased initially from the intact state following implantation of the artificial disc and again after the first foraminotomy, however, neither of these increases was significant. Subsequent foraminotomies following the initial foraminotomy caused only a minimal change in segmental motion. This indicates that any destabilizing effect from the foraminotomy occurred after violation of the initial joint and was not affected by additional foraminotomies. The biomechanical results of this analysis illustrate that a posterior foraminotomy can be performed in the setting of a pre-existing CDA without causing a significant increase in segmental range of motion.

CDA has had clinical and biomechanical success in recent literature. In spite of this, recurrent/persistent symptoms following CDA, as in ACDF, are inevitable in some patients and are one of the more common reasons for surgical revision. The ideal approach to revision surgery in the setting of CDA is unknown. One option entails a repeat anterior approach with explant of the arthroplasty device and conversion to a traditional ACDF. The drawbacks of this approach include higher complications with the repeat approach<sup>16,17</sup> and loss of any advantages seen with preservation of segmental motion. A potentially more attractive alternative would be to perform the revision decompression from a posterior approach via laminoforaminotomy. This would obviate the need to dissect scar tissue in the anterior neck and allow preservation of the arthroplasty device. However, the safety of posterior foraminotomy in the setting of CDA is largely unknown.

The potentially destabilizing effects of posterior laminoforaminotomy have been known for decades. In the early 1990s, Zdeblick, et al.<sup>18,19</sup> published biomechanical results on the consequences of posterior foraminotomy, stating that greater than 50% resection of the facet joint and capsule created segmental instability. More recent studies have reported clinical data regarding iatrogenic instability resulting from posterior laminoforaminotomy. In 2009, Jaganathan, et al.<sup>13</sup> published loss of cervical lordosis in 20% and instability in 4.9% of foraminotomy patients. Additionally, Clarke, et al.<sup>20</sup> cited a 6.7% risk of adjacent segment disease at 10 years following posterior foraminotomy. While there are downsides, when performed correctly, the procedure is safe and highly effective for the treatment of cervical radiculopathy.<sup>11,12</sup>

There exists only limited data regarding the combination of CDA and posterior foraminotomy. A retrospective clinical series published by Sekhon, et al.<sup>21</sup> examined the results of CDA in patients who had undergone a prior posterior foraminotomy. They reported favorable clinical outcomes with no cases of instability resulting from the addition of a CDA to a cervical spine with a foraminotomy. A biomechanical analysis by Roberto, et al.<sup>22</sup> in 2010 examined the effects of uncinete resection, posterior longitudinal ligament excision, foraminotomy, and laminectomy on cervical ROM and translation following ProDisc C implantation. The authors found some statistically significant increases in ROM from the intact specimens to the foraminotomy specimens, but concluded that the posterior decompressive procedures were safe in the setting of cervical disc arthroplasty. A subsequent biomechanical study in 2010 by Kikkawa, et al.<sup>23</sup> studied the effects of laminoplasty in combination with cervical disc arthroplasty. Similarly, the authors reported no significant ROM increases when laminoplasty was combined with CDA. This current study aimed to focus primarily on the effects of posterior foraminotomy in the setting of CDA and our results echo that of the previously mentioned biomechanical reports. The flexion/extension (sagittal motion) plane was most affected by posterior

decompressive procedures, but the addition of a posterior foraminotomy did not significantly increase segmental ROM.

There are several limitations to this study. The specimens included for analysis are from elderly people who inevitably had some degeneration of their cervical spines. This fact makes the studied population different than the usual CDA candidates, namely younger patients with limited disc degeneration and radiculopathy. However, this can potentially be viewed as a strength of the study. Performing wide discectomies and decompressions in specimens with some degree of spondylosis likely causes more segmental destabilization than the same procedure would in a young, non-degenerative spine. Secondary to this, the ROM differences found in our analysis may be overestimated from what would be observed in younger patients who are more likely to undergo CDA. Another weakness is that this method of biomechanical testing allows for ROM analysis at only one point in time. Therefore, it is difficult to draw conclusions about the development of segmental instability over time.

A question that remains when applying this biomechanical data to clinical use is determining whether a statistically significant difference, or lack of, represents one that is clinically significant. The addition of a foraminotomy to spines with a preexisting CDA caused a 2.6° (18%) increase in sagittal plane (flexion/extension) motion. While this was not statistically significant, it is unknown whether this ROM increase would manifest clinically.

CAD, to date, has been clinically successful and offers several theoretical advantages over traditional fusion procedures. As the number of people living with a CDA continues to increase, so will the need for surgical revision. In the setting of recurrent/persistent radiculopathy, posterior foraminotomy obviates the need to re-expose the anterior spine and offers the advantage of being able to leave the CDA intact. Iatrogenically induced instability from this approach is a valid concern. However, the results of this cadaveric analysis indicate biomechanically that when less than 50% of the facet joint is removed, bilateral and even supra-adjacent level foraminotomies can be performed without significantly increasing segmental ROM.

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## Risk Factors for Infection and Amputation following Open, Combat-Related Calcaneus Fractures

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### INTRODUCTION

Open calcaneus fractures represent a severe and often disabling injury, occurring in less than 8.5% of all calcaneus fractures,<sup>1</sup> with subsequent complications occurring in up to 78% of cases.<sup>1-5</sup> The severity of the injury, soft tissue compromise, fracture displacement, and fracture comminution present unique challenges in the treatment of open calcaneus fractures. These injuries are often complicated by non-anatomic reduction, limited subtalar motion, wound necrosis, infection, guarded functional outcomes, injury to passing neurovascular and tendinous structures, and frequent eventual amputation.<sup>1-9</sup>

The natural history of open calcaneus fractures is poorly understood and the variables that allow for successful limb salvage have not been described. Likewise, few studies have assessed treatment and outcomes of open calcaneus fractures. The purpose of this study was to further characterize high-energy open calcaneus fractures by identifying risk factors that may predict infection, amputation, and poor outcomes. We hypothesized that more severe soft tissue injury, as measured by both the Gustilo and Anderson classification and the size and location of the wound(s),<sup>4</sup> would be the strongest predictor of infection and amputation.

### METHODS

After institutional review board approval, all patients receiving treatment for an open calcaneus fracture sustained between March 2003 to August 2010 during Operation Iraqi Freedom (OIF) and Operation Enduring Freedom (OEF) were identified from the Armed Forces Health Longitudinal Technology Application (AHLTA), the Joint Theater Trauma Registry (JTTR), and the local Surgical Scheduling System (S3). The JTTR is a database of medical treatment information on patients in a theater of combat operations treated at U.S. military medical facilities. AHLTA is the electronic medical record and coding system for the military healthcare system. The S3 is an electronic operative case log of all surgical procedures at our facility. All databases were queried to identify patients with an open or closed calcaneus fracture (ICD-9 code 825.0, CPT codes 28400, 28415, 28420, 28406).

Inclusion criteria were U.S. service members who sustained an open calcaneus fracture while performing combat operations and who subsequently received treatment at a single tertiary referral hospital. Patients who received an immediate amputation within the first 24 hours of injury were excluded. Thus, all patients in this study population were deemed to have severe lower extremity trauma resulting in an open

calcaneus fracture, but the injury did not require immediate amputation for a grossly unsalvageable limb or to preserve life.

Abstracted data included patient demographics, mechanism of injury, wound size and location, Gustilo and Anderson<sup>10,11</sup> and Sanders<sup>12,13</sup> fracture classifications, interval and definitive treatment procedures, adjunctive procedures (rotational or free tissue transfer, skin graft, and neurovascular procedures), as well as ipsilateral and contralateral orthopaedic injuries. The definitive treatment was defined as the procedure after injury which, after healing, would allow reduced fracture union and eventual weight bearing. The reported complications were documented for deep and superficial infection, need for late amputation, and type and timing of revision surgery. Functional outcomes were evaluated by visual analog pain scale and recreational status as assessed by Tegner activity level.<sup>14</sup>

### RESULTS

From March 2003 to July 2010, 583 U.S. military personnel sustained a calcaneus fracture during OEF or OIF. Two hundred eighty-three individuals received treatment at one of our institutions for a combat-related calcaneus fracture and 122 patients sustained open fractures. Thirty-three patients were transferred to another hospital during their initial hospital admission and were thus excluded from the study population due to lack of in-patient data. Complete records were available for 89 patients who sustained 102 combat-related open calcaneus fractures and who were followed for a mean of four years (range, 5 to 92 months). The injury and treatment characteristics are presented in Table 1.

#### Limb Salvage versus Amputation

During the study period, 42% (43/102) of open calcaneus fractures were treated with an amputation (42 transtibial, 1 knee disarticulation). Extremities that failed limb salvage were more likely to have an associated ipsilateral forefoot fracture ( $p < 0.0001$ ), talus fracture ( $p < 0.0001$ ), plantar wound ( $p < 0.0001$ ), and culture positive wound infection ( $p < 0.0001$ ). Wound size was significantly associated with amputation ( $p < 0.0001$ ; Table 2). The mean total size of salvaged open calcaneus fracture wounds was  $15 \text{ cm}^2 \pm 13 \text{ cm}^2$  versus  $51 \text{ cm}^2 \pm 58 \text{ cm}^2$  ( $p < 0.0001$ ) in the amputation cohort. Gustilo and Anderson fracture type was significantly associated with amputation, with type I (0%), type II (8%), type IIIA (29%), type IIIB (55%), and type IIIC (100%) ( $p < 0.0001$ ). Ipsilateral tibia fractures, spine fractures, and the absence of plantar sensation at presentation were not significantly associated with eventual amputation. In a multivariate Cox Proportional Hazards Survival model with amputation as the end point, blast mechanism of injury, location and larger size of open wound in  $\text{cm}^2$ , and escalating Sanders and Gustilo and Anderson classification types (all  $p < 0.05$ ) were predictive of eventual amputation. At final follow-up, patients who

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**Table 1. Demographic and Injury Characteristics of Open Calcaneus Fractures Treated at Walter Reed National Military Medical Center from 2003 - 2010**

Characteristic	No. (range or %)
Total open calcaneus fractures	102
Median age (range) (yr)	26 (19-44)
Injury severity score (range)	18.6 (5-57)
Sex*	
Male	85 (96)
Female	4 (4)
Method of injury	
Improvised explosive device	75 (74)
Gun shot	8 (8)
Rocket-propelled grenade	16 (16)
Fall from height	1 (1)
Vehicular accident	2 (2)
Associated major ipsilateral lower extremity fractures	
Femur	7(7)
Tibia	44 (43)
Talus	49 (48)
Method of definitive fixation	
Plate-and-screw fixation	27 (26)
Circular external fixation	11 (11)
Pin or screw fixation	16 (16)
Nonoperative	14 (14)
Subtalar arthrodesis	12 (12)
Amputation before fixation attempt	22 (22)
Wound location	
Plantar	55 (54)
Medial	9 (9)
Lateral	10(10)
Dorsal	16 (16)
Posterior	12 (12)
Wound size (cm <sup>2</sup> ) (mean 30 cm <sup>2</sup> )	
≤ 20	65 (64)
21 to 39	19 (19)
≥ 40	18 (18)
Gustillo and Anderson classification	
I	3 (3)
II	12 (12)
IIIA	31 (30)
IIIB	49 (48)
IIIC	6 (6)
Plantar Sensation	
Intact	36 (35)
Diminished	55 (54)
Absent	11 (11)
Common culture positive deep infections (n = 47)	47 (46)
<i>Acinetobacter</i>	21 (45)
<i>Staphylococcus</i>	12 (26)
<i>Pseudomonas</i>	7 (15)
<i>Enterococcus</i>	11 (23)
<i>Klebsiella</i>	6 (13)
Mold species	2 (4)
<i>Enterobacter</i>	3 (6)
Other	2 (4)
Multiple organisms	17 (36)
Soft tissue reconstruction (n = 28)	
Skin graft	22 (22)
Free tissue transfer	6 (6)

Total procedures	13.7 (2-47)
Total procedures after definitive treatment	3.3 (0-14)
Amputation (n = 43)	
Time from injury to amputation (days)	120 (1-647)
Transtibial	42 (98)
Through knee	1 (2)
Outcomes (n = 100)	
Return to duty (n = 100)	38 (38)
Medical board	62 (62)
Visual analog scale pain	3.2 (0-7)
Tegner activity level	4.3 (0-7)

\* The values are given as the number and the percentage or range in parentheses.

had undergone amputation had lower visual analogue scale scores for pain (2.1 versus 4.0,  $p < 0.0001$ ) and higher Tegner activity levels (5.4 versus 3.5,  $p < 0.0001$ ) than limb salvage patients.

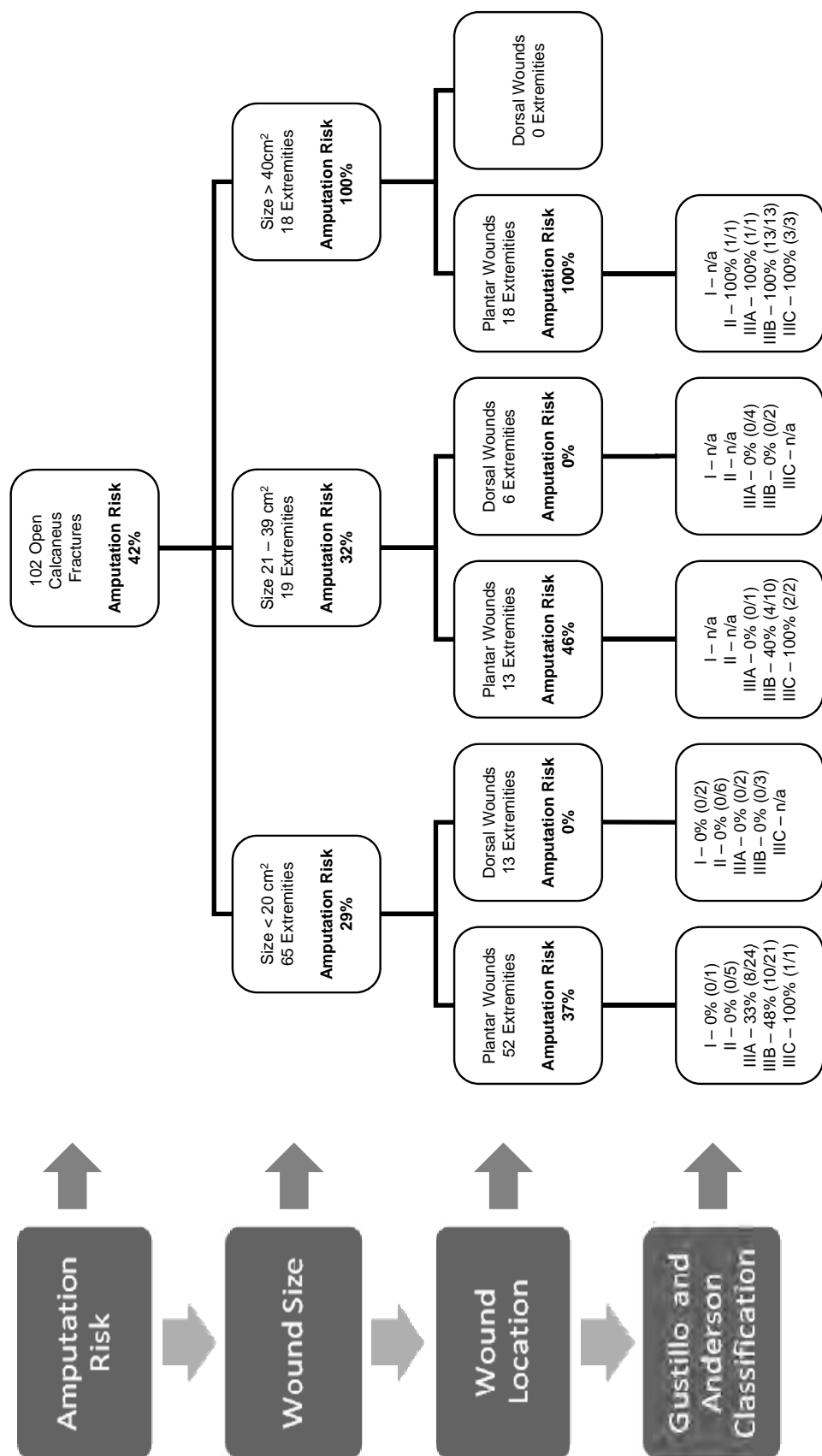
Complications following limb salvage and amputation were common. Compared to extremities that received early amputation, the group that received a delayed amputation required more total operative procedures during the study period (17 versus 11,  $p = 0.001$ ). Sixty-six percent (29/43) of patients with an amputation required a re-operation and/or revision procedure. The most common revision procedures for those who underwent amputation were irrigation and debridement (63%; 27/43), revision wound closure (49%; 21/43), and hardware removal (19%; 8/43). Comparatively, 83% (49/59) of patients who continued limb salvage treatment required a revision surgery. The most common revision procedures in the limb salvage group were implant removal (42%; 25/59), exostectomy (37%, 22/59), and revision wound closures (34%; 20/59).

### Early versus Delayed Amputation

A late amputation (> 12 weeks from injury) was performed in 15% (15/102) of open calcaneus fractures compared to 27% (28/102) undergoing early amputation (24 hours to 12 weeks from injury). The mean time to early amputation was 19 days (range, 1 to 53) versus 312 days (range, 84 to 647) in the delayed amputation group. All patients in the late amputation cohort received a transtibial amputation; one knee disarticulation was performed in the early period. Patients receiving delayed or late amputation did not demonstrate significantly different injury severity score, Gustilo and Anderson fracture severity, wound size, wound location, presence of plantar sensation, need for tissue transfer, associated fracture patterns, or final Tegner activity level. Patients receiving a late amputation underwent statistically more revision surgeries (7.2 versus 1.9,  $p < 0.001$ ) and were more likely to receive definitive fracture treatment in a circular external fixator ( $p < 0.001$ ).

### DISCUSSION

Open calcaneus fractures account for 8.5% of all calcaneus fractures in the civilian trauma setting.<sup>1</sup> Presently, 43% of wounded warriors with calcaneus fractures present to our institution more than 24 hours from injury with a combat-related open calcaneus fracture and a salvaged lower extremity. Open calcaneus fractures are burdened by high infection rates, compromised soft tissues, high malunion and nonunion rates, substantial patient morbidity, and high rates of concomitant inju-



**Figure 1.** Risk factors for amputation follow open calcaneus fractures. Overall, 42% of patients with an open calcaneus fracture required amputation. All Gustillo and Anderson IIIC fractures required amputation. Gustillo and Anderson IIIB fractures with plantar wounds required amputation in 61% of extremities, regardless of the size of the wound.



**Table 2. Risk Factors for Amputation and Infection following Open Calcaneus Fracture**

Characteristic	No.	Amputation (%)	Likelihood ratio	p value	Infection	Likelihood ratio	p value
Wound size (cm <sup>2</sup> ) (mean 30 cm <sup>2</sup> )							
≤ 20	65	19 (29%)			28 (43%)		
21 to 39	19	6 (32%)			10 (52%)		
≥ 40	18	18 (100%)	36.6	< 0.0001	9 (50%)		0.107
Ipsilateral lower extremity fractures							
Femur	7	5 (71%)		0.104	4 (57%)		0.543
Tibia	44	23 (53%)		0.072	23 (53%)		0.274
Talus	50	29 (58%)	10.3	0.001	30 (60%)	7.7	0.006
Forefoot	67	37 (55%)	14.7	< 0.0001	37 (55%)	6.7	0.01
Method of definitive fixation							
Plate-and-screw fixation	27	4 (15%)			9 (33%)		
Circular external fixation	11	5 (45%)			6 (55%)		
Pin or screw fixation	16	6 (38%)			9 (56%)		
Nonoperative	14	--			3 (21%)		
Subtalar arthrodesis	12	6 (50%)		0.086	6 (50%)		
Amputation before fixation attempt	22	--			14 (64%)	9.3	0.11
Wound location							
Dorsal	19	0 (0%)			5 (26%)		
Plantar	83	43 (52%)	23.9	< 0.0001	42 (51%)	3.8	0.055
Gustilo and Anderson classification							
I	3	0 (0%)			1 (33%)		
II	12	1 (8%)			3 (25%)		
IIIA	32	9 (28%)			10 (31%)		
IIIB	49	27 (55%)			29 (59%)		
IIIC	6	6 (100%)	27.2	< 0.0001	4 (67%)	12.2	0.042
Plantar sensation							
Intact	36	11 (31%)			14		
Diminished	55	25 (45%)			27		
Absent	11	7 (64%)	4.4	0.116	6	1.27	0.531
Culture positive deep infection	47	28 (60%)	11.1	0.001	--	--	--

Amputation was significantly associated with wounds ≥ 40 cm, ipsilateral talus and forefoot fractures, initial definitive fixation using subtalar arthrodesis, plantar wounds, and culture positive deep wound infections. Number of patients are documented with percentage in parentheses. Likelihood ratios are provided for significant variables.

ries, often necessitating multiple revision surgeries or delayed amputation. To date, small series have reported complications following open calcaneus fractures in up to 78% of injuries.<sup>1-5,7</sup> These studies report soft tissue complications in 44% of open calcaneus fractures<sup>4</sup>, deep tissue infection in 10% to 39%,<sup>1-4</sup> and eventual amputation in 0 to 14%.<sup>1-4,9</sup> The factors that predict infection, amputation, and poor outcomes remain ill-defined. The purpose of this study was to characterize high-energy open calcaneus fractures by identifying risk factors that may predict infection and amputation. We hypothesized that more severe soft tissue injury, as measured by the Gustilo and Anderson classification and size of the wound<sup>4</sup> would be the strongest predictor of infection and amputation.

The incidence of amputation and risk factors that predict amputation following open calcaneus fractures had not been previously identified.<sup>1-4,15</sup> No prior study has demonstrated a significant correlation between Gustilo and Anderson fracture type and ampu-

tation following an open calcaneus fracture.<sup>1-5,8,16</sup> Heir, et al.<sup>1</sup> found that a subset of patients with Gustilo and Anderson type IIIB fractures and wounds in a non-medial location were more likely to require amputation. To our knowledge there have been a total of 78 type III open calcaneus fractures reported in the English literature, with a 22% incidence of amputation following an initial period of limb salvage in this population (n = 17; range, 0 to 33%).<sup>1-5,8,9,16</sup> Our study demonstrated that increasing Gustilo and Anderson fracture type is an independent risk factor for amputation following open calcaneus fractures, irrespective of the location of the predominant wound. We note an amputation rate of 47% in 87 patients with type III open calcaneus fractures, which is notably higher than previously reported series.<sup>1-5,8,16</sup> The larger soft tissue deficits and more frequent associated injury patterns that result from blast trauma likely contribute to the higher amputation rate in this study.

The characterization of injuries leading to early versus delayed amputation is infrequently reported<sup>17-23</sup> and there are no studies which specifically evaluate the impact of open calcaneus fractures on the timing of amputation. Additionally, given the infrequent nature of open calcaneus fractures and the limited series evaluating the natural history of this injury, inconsistency exists regarding the definition of amputation timing (early or late) and the rate of amputation. In our series, the overall amputation rate was 42%, with 15% of open calcaneus fractures eventually requiring a late amputation more than 12 weeks after injury. These findings are consistent with a recent evaluation of amputations in the current military conflicts, in which the overall late amputation rate for all amputees was 15%.<sup>17</sup> This is in contradistinction to the lower extremity assessment project (LEAP) study group, which found that after a two-year follow-up, late amputations, defined as occurring after the initial hospitalization, occurred in 3.9% of cases.<sup>17</sup> It is possible that the more severe soft tissue injuries and multiple associated fractures may account for this difference. It is also conceivable that in our military practice environment, in an open access healthcare system with a sophisticated amputee rehabilitation center, it is more socially acceptable and functionally practicable for an injured young service member to pursue amputation.

Outcomes following open calcaneus fractures have demonstrated poor functional outcomes in Gustilo and Anderson type III fractures,<sup>4</sup> while more satisfactory results have been reported in less severe open fracture wounds,<sup>1,3,9</sup> wounds in a medial location,<sup>3,16</sup> and after staged treatment that first addresses the soft tissue wound.<sup>1,4,8,16</sup> In this series of severe open calcaneus fractures, patients who pursued limb salvage were able to perform light labor and had significantly higher amounts of pain compared to patients who received eventual amputation. After two-year<sup>15</sup> and seven-year<sup>6</sup> follow-up, the LEAP study group found no difference in functional outcomes between those patients who underwent limb salvage or amputation using the Sickness Impact Profile. Functional outcomes were poor in both groups, worse after longer follow-up, and not affected by the severity of the fracture or soft tissue injury. Our results demonstrate significantly greater Tegner activity levels and lower pain scores in patients who received an amputation. These differences may be attributed to the younger age, centralized rehabilitation, and increased amputee resources available for our patient population. Long-term outcomes of amputation in our young patient population are not known; while early outcomes demonstrate improved function, we believe that amputation should not be performed without careful deliberation with the patient and on an individualized basis. Additionally, we found no difference in activity level or pain in amputations performed in the early versus the late period. This is in contrast to a recent LEAP study, which reported more complications in patients undergoing late amputation for any lower extremity trauma.<sup>23</sup> The initial pursuit of limb salvage followed by late amputation, while requiring additional procedures and a longer treatment course in our study, does not appear to influence ultimate Tegner activity level or pain.

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# The Ventral Lamina and Superior Facet Rule: a Morphometric Analysis for Ideal Thoracic Pedicle Screw Starting Point

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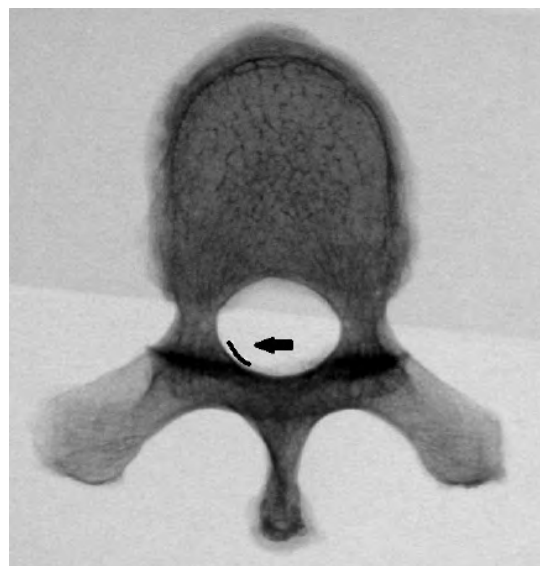
## INTRODUCTION

Pedicle screw instrumentation has become increasingly popular for fixation of the thoracic spine for degenerative, deformity, tumor, and trauma pathologies. Given the inherent dangers associated with pedicle screw placement, the topographic anatomy has been extensively detailed and described in the literature.<sup>1-7</sup> This has allowed surgeons to carefully find the appropriate anatomic starting point, as well as to determine the optimal screw length and trajectory. Despite the recent plethora of literature detailing the anatomic starting points, the safety and efficacy of screw insertion, and the expanding scope of practice utilizing thoracic pedicle screws, there are no objective, definable landmarks to assist with free-hand insertion of pedicle screws. The slightest misunderstanding or error can lead to an unwanted cortical breach, resulting in potential neurologic complication, and the risk of iatrogenic injury from an erroneously placed screw is more catastrophic in the thoracic spine versus the lumbar spine.

With our own increasing surgical experience, we have noted a reproducible and unique anatomic structure known as the “ventral lamina.” The ventral lamina is formed by the roof of the spinal canal and is confluent laterally with the medial pedicle wall (Figure 1). An increased understanding of its three-dimensional structure and relationships has significantly improved our ability to place screws safely in pedicles previously thought impossible to instrument (*i.e.*, hypoplastic, deformed, or those with small cancellous channels). This cadaveric study is designed to test the hypothesis that the lateral border of the ventral lamina, where it is contiguous with the medial pedicle wall, occurs medial to the midline of the superior articular facet, thereby serving as a reproducible anatomic guide for pedicle screw instrumentation of the thoracic spine. We set out to 1) determine the true morphometry of the ventral lamina, and 2) to establish a reproducible, ideal medial-lateral thoracic pedicle screw starting point based on the anatomic relationship of the ventral lamina and pedicle with the superior articular facet (SAF), which we term the “superior facet rule.”

## METHODS

One-hundred fifteen thoracic spine vertebral levels were individually disarticulated and divested of all soft tissue, ensuring the



**Figure 1.** Axial radiograph of thoracic vertebra, demonstrates the ventral lamina (arrow) and its confluence with the medial pedicle wall.

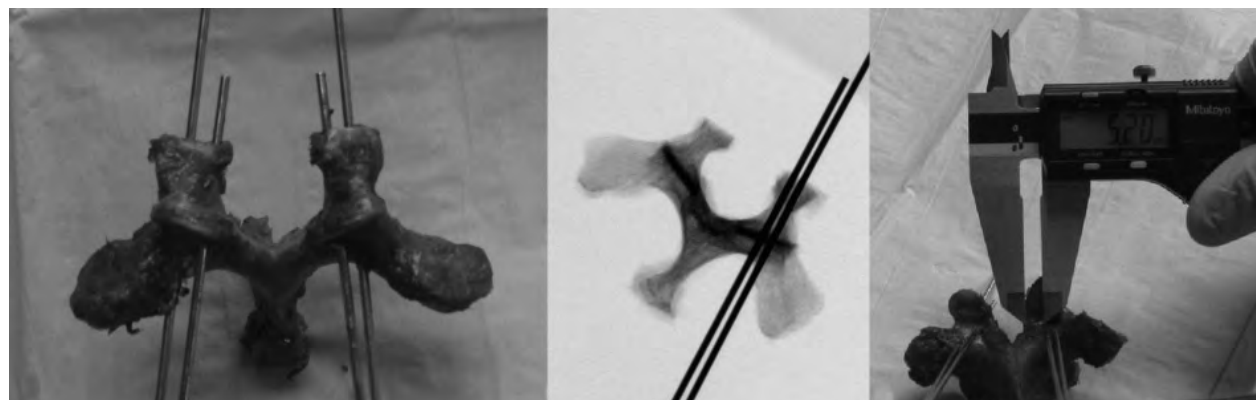
osseous surface of each SAF, lamina, pedicle, and spinous process were clearly visualized. Each individual level was then prepared by removing the vertebral body with a coronal saw cut at the junction of the pedicle and body. Kirschner wires were then placed in a retrograde fashion along the four boundaries of the pedicle canal (medial, lateral, caudad, and cephalad) (Figure 2). Using digital calipers, the following distances were measured in millimeters (mm):

- 1) width of the SAF at longitudinal mid-point
- 2) medial border of SAF to medial pedicle wall (MPW)
- 3) midline of vertebra (cephalad ridge of spinous process) to MPW
- 4) lateral border of SAF to lateral pedicle wall
- 5) pedicle diameter at the isthmus

Measurements were analyzed to calculate the anatomic relationship of the ventral lamina and center of the pedicle (COP) to the midline of the SAF. Data analysis was performed using mean  $\pm$  standard deviation and 95% confidence interval. This study was performed using intramural institutional funding; there are no potential conflicts of interest directly related to this study.

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**Figure 2.** Image of test specimen with vertebral body removed, and K-wires placed along medial and lateral boundaries of pedicle (A); axial radiograph of test specimens with K-wires placed (B); image of digital caliper measurement (C).

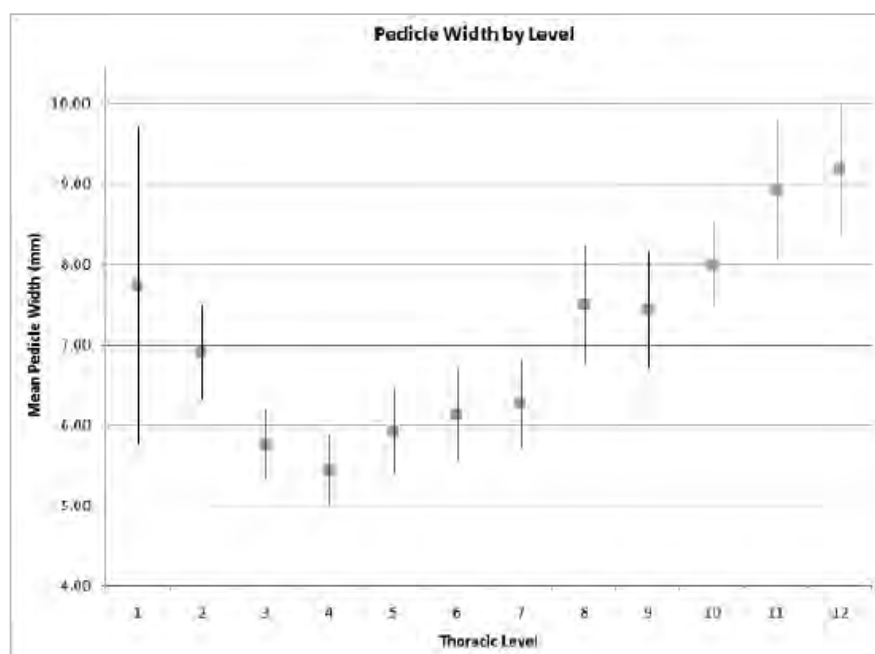
## RESULTS

Two hundred twenty-nine pedicles were measured. One pedicle was excluded due to fracture of the SAF during disarticulation. The ventral lamina was clearly identifiable at all levels, and in every specimen, forming the roof of the spinal canal and confluent with the medial pedicle wall. The smallest average pedicle width measurements occurred at the T3 to T7 levels (range, 5.45 mm to 6.27 mm), with the largest average measurements occurring at T11 and T12 (range, 8.94 mm to 9.19 mm). This is similar to previous studies evaluating pedicle morphometry, which showed the smallest pedicle isthmic diameters

occurring in the mid-thoracic region between T4 and T7.<sup>8</sup> The average pedicle width for all pedicles was  $7.06 \text{ mm} \pm 1.86 \text{ mm}$  (CI = 6.82 mm to 7.30 mm) (Chart 1, Table 1).

The mean distance from the midline of the SAF to the ventral lamina was  $1.36 \text{ mm} \pm 1.23 \text{ mm}$  medial (CI = 1.19 to 1.52). The ventral lamina was lateral to the midline of the SAF in 34 (14.85%) pedicles, with a mean distance of only  $0.52 \pm 0.51 \text{ mm}$  lateral (CI = 0.35 to 0.69). There was only 1 (0.43%) ventral lamina that was  $> 2 \text{ mm}$  lateral, and it was found to be 2.18 mm lateral (Chart 2, Table 2).

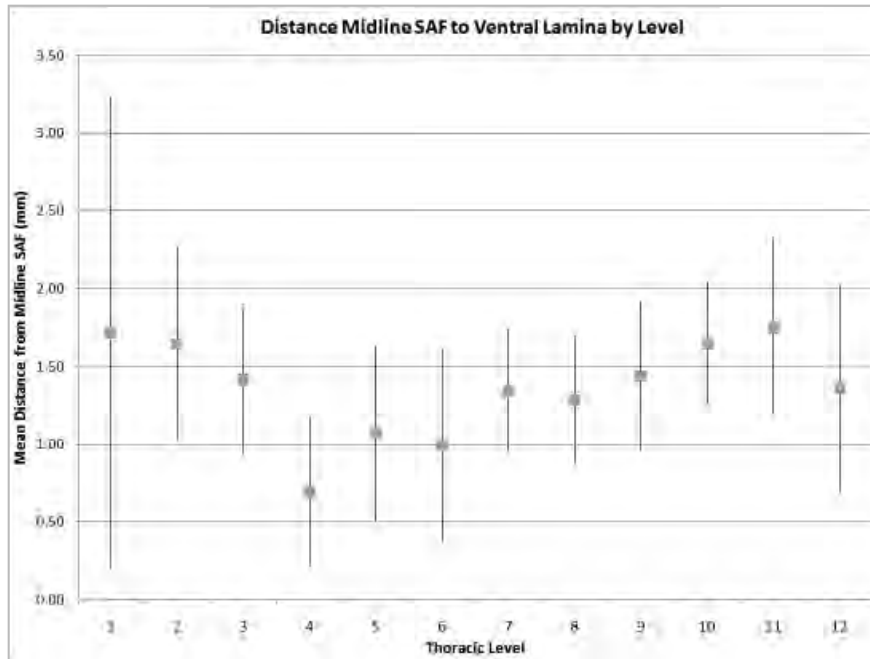
The mean distance from the midline of the SAF to the center of the pedicle (COP) was  $2.17 \text{ mm} \pm 1.38 \text{ mm}$  lateral (CI = 1.99 to



**Chart 1.** Mean pedicle width by level (mm).

**Table 1. Average Pedicle Width per Level (mm)**

	N =	Mean	Standard deviation	CI
T1	9	7.73	3.03	1.98
T2	14	6.91	1.11	0.58
T3	24	5.77	1.08	0.43
T4	20	5.45	1.00	0.44
T5	20	5.94	1.20	0.53
T6	20	6.13	1.30	0.57
T7	20	6.27	1.25	0.55
T8	20	7.51	1.67	0.73
T9	22	7.44	1.67	0.73
T10	20	8.01	1.24	0.54
T11	20	8.94	2.02	0.88
T12	20	9.19	1.86	0.82
<b>TOTAL</b>	<b>229</b>	<b>7.06</b>	<b>1.86</b>	<b>0.24</b>

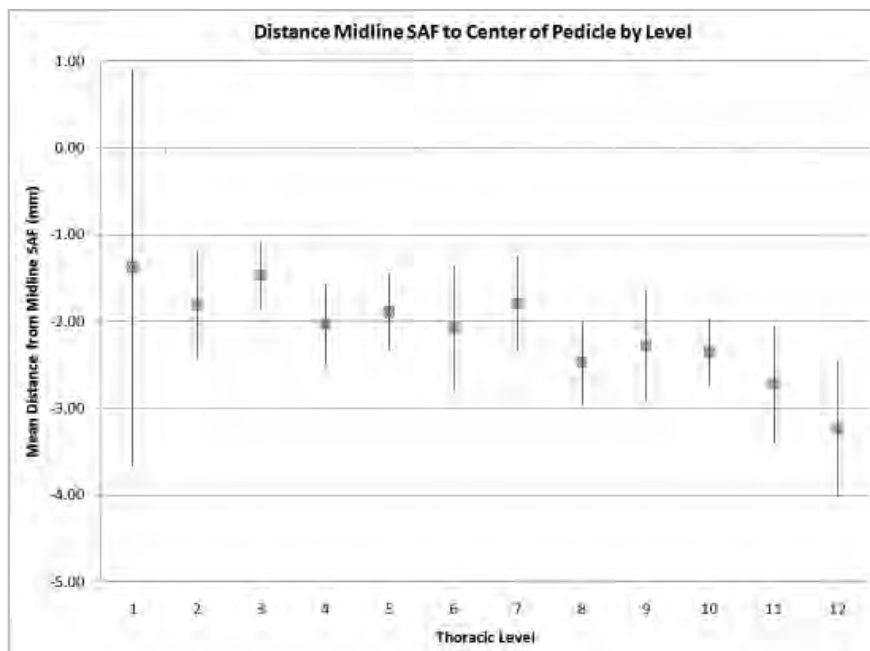


**Chart 2.** Mean distance midline SAF to ventral lamina by level (mm).

<b>Table 2. Mean Distance from Midline Superior Articular Facet to Ventral Lamina (mm)</b>				
	N =	Mean	Standard deviation	CI
T1	9	1.72	2.32	1.52
T2	14	1.65	1.17	0.62
T3	24	1.42	1.20	0.48
T4	20	0.70	1.10	0.48
T5	20	1.08	1.28	0.56
T6	20	0.99	1.42	0.62
T7	20	1.34	0.90	0.39
T8	20	1.29	0.95	0.42
T9	22	1.44	1.08	0.47
T10	20	1.65	0.89	0.39
T11	20	1.75	1.33	0.58
T12	20	1.36	1.51	0.66
<b>TOTAL</b>	<b>229</b>	<b>1.36</b>	<b>1.23</b>	<b>0.16</b>

2.35). The COP was medial to the midline of the SAF in 11 (4.80%) pedicles, with a mean distance of only  $0.76 \text{ mm} \pm 0.76 \text{ mm}$  medial. The most medial COP was 2.32 mm medial to the midline of the

SAF (Chart 3, Table 3). We also found the COP to be  $13.17 \text{ mm} \pm 2.47 \text{ mm}$  caudad (CI = 12.85 to 13.49) to the superior border of the SAF.



**Chart 3.** Mean distance midline SAF to center of pedicle by level (mm).

<b>Table 3. Mean Distance from Midline Superior Articular Facet to Center of Pedicle (mm)</b>				
	N =	Mean	Standard deviation	CI
T1	9	-1.37	3.50	2.29
T2	14	-1.81	1.18	0.62
T3	24	-1.47	0.97	0.39
T4	20	-2.03	1.06	0.47
T5	20	-1.89	1.03	0.45
T6	20	-2.07	1.64	0.72
T7	20	-1.79	1.24	0.54
T8	20	-2.47	1.11	0.49
T9	22	-2.28	1.47	0.64
T10	20	-2.35	0.88	0.39
T11	20	-2.72	1.54	0.68
T12	20	-3.23	1.79	0.78
<b>TOTAL</b>	<b>229</b>	<b>-2.17</b>	<b>1.38</b>	<b>0.18</b>

In addition, we evaluated all calculated distances, irrespective of vertebral level, of the ventral lamina and the COP (Figure 4) in relation to the midline SAF ordered in increasing distance from the midline SAF. We found a parabolic relationship (Figure 4) and this was used to determine the ideal anatomic starting point approximately 2 mm to 3 mm lateral to the midline SAF.

## DISCUSSION

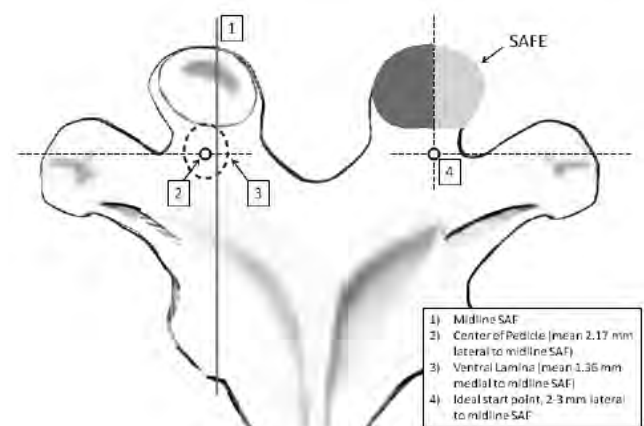
Given the individual variability in pedicle anatomy, as well as differences in surgeon training/experience with thoracic pedicle screw placement, the primary purpose of this study was to verify our observation regarding the presence of the ventral lamina as a reproducible, anatomic structure that can serve as an anatomic “map” for the ideal medial-lateral placement of thoracic pedicle screws. In 85% of our studied specimens, the ventral lamina was found to be medial to the midline of the SAF. In the remaining 15%, the ventral lamina was lateral to the midline of the SAF by a mean distance of 0.52 mm, and only one (0.43%) ventral lamina was > 2 mm lateral to the midline of the SAF. This statistic alone further solidifies the tenet that if the surgeon begins the starting point 2 mm lateral to the midline of the SAF, there is a 0.43% chance the starting point will be medial to the pedicle or into the spinal canal. The center of the pedicle was also found to be a mean distance of 2.17 mm lateral to the midline of the SAF, and was lateral to midline in 95% of pedicles. Therefore, results from our morphometric study suggest the ideal starting point for thoracic pedicle screw insertion should be 2 mm to 3 mm lateral to the midline of the SAF. We term this reproducible, anatomic landmark the “superior facet rule” (Figure 3). This consistent anatomic relationship can be used as a reliable guide for thoracic pedicle screw insertion in the center of the pedicle, and to avoid penetration into the spinal canal without having to adjust the starting point depending on the thoracic vertebral level being instrumented. The surgeon should also take into consideration the fact that the medial pedicle wall is two to three times thicker than the lateral pedicle wall.<sup>1</sup> The medial pedicle wall can be envisioned as a “buttress,” serving as a boundary for a “funnel,” or path, for the trajectory of the pedicle screw.

Previous studies have recommended various anatomic landmarks for appropriate pedicle screw placement; however, no standard, objective anatomic landmark for thoracic pedicle screw insertion exists.<sup>9-14</sup> Cinotti, et al.<sup>9</sup> also used the superior facet as an anatomic landmark for thoracic pedicle screw insertion and found the pedicle axis intersecting the middle of the SAF in 15% of specimens, lateral two-thirds of the SAF in 62%, and lateral border of the SAF in 23%. The authors found the COP to be at or lateral to the midline SAF in 100% of specimens, which is similar to our findings (95%) concerning the lateral relationship of the COP to the midline SAF. In the same study, an entry point at the midline SAF resulted in a 10% rate of pedicle cortex violation, compared to no pedicle cortex violation using the entry point along the lateral two-thirds of the SAF.<sup>9</sup> Therefore, the authors recommended a starting point at the intersection between the superior border of the transverse process and the lateral two-thirds of the SAF.

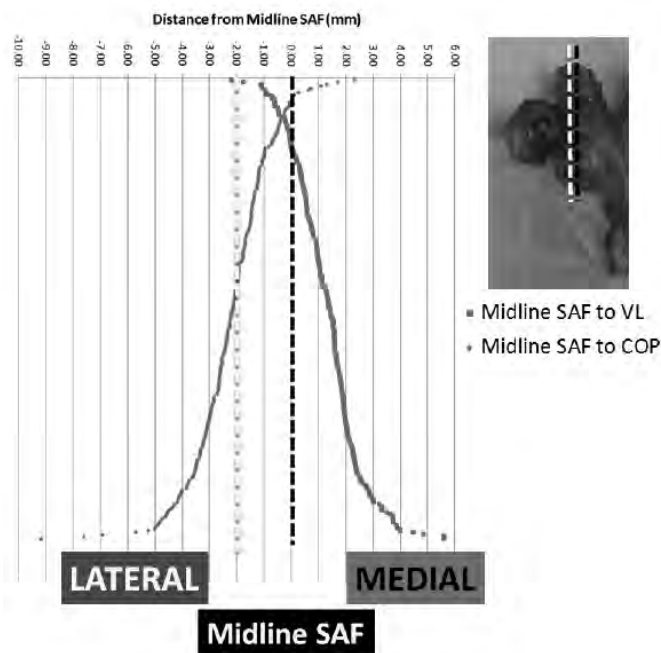
Recently, Yang, et al.<sup>14</sup> described the utility of the nutrient foramen of the lamina as a guide to placement of pedicle screws in the thoracic spine. However, the nutrient foramen was not found uniformly at all thoracic levels, and was found in only 63% of specimens. This lack of reproducibility may make pedicle screw insertion difficult if one relies on this anatomic landmark alone.<sup>14</sup> This is in comparison to our current study, which found the superior facet rule to be 95% reproduc-

ible and consistent at nearly all thoracic levels. In another study, Kim, et al.<sup>15</sup> describe the ideal pedicle screw starting point in the thoracic spine. However, the authors describe a dynamic and changing starting point depending on the thoracic level. For the lower thoracic regions, the starting point is described as the junction of the bisected transverse process and the lamina at, or just medial to, the lateral aspect of the pars. As instrumentation progresses cephalad to the apical mid-thoracic region, the starting point trends toward a more medial and cephalad starting point. At T7 to T9, the anatomic starting point is described as the confluence of the transverse process, lamina, and the superior articular facet. Finally, at the most cephalad levels, the starting point then changes again to progress more lateral and caudad, described as the junction of the bisected transverse process and the lamina at the lateral pars. Using this technique, the authors reported only a 1.7% incidence of medial cortical breach with no neurologic complications in a total of 577 screws.<sup>15</sup> Despite the safety profile highlighted by use of the free-hand technique as described by Kim, et al.,<sup>15</sup> of particular mention is the fact that the success of the free-hand technique used in this study was achieved by a surgeon with clinical experience in placing over 30,000 pedicle screws.

The difference between the thoracic and lumbar spine pedicle anatomy cannot be over emphasized, and makes the task of placing pedicle screws much more challenging in thoracic vertebrae. This is especially true within the mid-thoracic spine (between T4 and T7), where previous morphologic studies have found pedicles become significantly smaller.<sup>7,9,16</sup> This finding was also demonstrated in our study where the pedicles were found to be much smaller between T3 and T7, compared to those at the most caudad levels (T10-T12) (Chart 1). Also, when evaluating the mean distance of the ventral lamina from the midline SAF, we found the T4 to T6 ventral laminae were closest to the midline SAF (Chart 2). However, when evaluating the mean distance of the COP from the midline SAF, the COP trended more lateral with more caudad thoracic levels (Chart 3). Interestingly, the T4 to T6 center of pedicle measurements were not the most lateral, despite having ventral laminae closest to the midline SAF, and this relationship may contribute to the mid-thoracic pedicles being the most difficult to instrument. These factors can make complete containment



**Figure 3.** Illustration of superior facet rule, with relationship of midline SAF (solid line) to ideal starting point (small circle), 2 mm to 3 mm lateral to midline SAF.



**Chart 4.** Combined measurements of all specimens, ordered in increasing distance from the midline SAF (black dashed line) irrespective of specimen or vertebral level, for the ventral lamina (squares) and center of pedicle (diamonds) (mm); the ideal medial-lateral starting point (2 mm lateral; light gray dashed line) is also depicted.

of pedicle screws extremely challenging, or sometimes even impossible in the mid-thoracic spine or at an apical thoracic vertebrae during deformity surgery, and misplaced screws or cortical breaches may be unavoidable. Some studies have suggested thoracic pedicles from T4 to T8 may not be wide enough for safe pedicle screw fixation.<sup>9,16</sup> However, these studies were performed with early generation instrumentation systems with the smallest available pedicle screw diameter of 5.5 mm. Our study demonstrated a mean inner pedicle isthmus diameter of 7.06 mm  $\pm$  1.86 mm (CI = 6.82 to 7.30), and the smallest measured pedicle isthmus was 5.45 mm. Medial cortical breach is of particular concern, given the proximity of the spinal cord to the medial cortical wall and the increased rate of neurologic injury reported with this error.<sup>7</sup> Despite the recognition of confounding residual imaging artifact (on post-operative computed tomography) as well as hypothesized “safe zones” for acceptable cortical violation, many situations still preclude the safe placement of pedicle screws.<sup>15,17-19</sup>

Several authors have demonstrated safety of free-hand pedicle screw placement in vivo, using only surface topography to guide screw placement.<sup>15,17,20-24</sup> Kim, et al.<sup>15</sup> demonstrated the free-hand placement of pedicle screws to be safe and effective, even in the setting of revision surgery;<sup>23</sup> however, misplaced screws still occur, and the resulting neurologic complications can be devastating.<sup>24</sup> Therefore, other authors have recommended the use of intra-operative fluoroscopy or image guidance techniques, which have been espoused to improve accuracy with screw insertion.<sup>18,19,25-30</sup> The accuracy rate of image-guided systems has been widely variable and very few in vivo studies actually exist. Several cadaveric studies have demonstrated an accuracy rate from as low as

80% to 99%.<sup>27-30</sup> A recent meta-analysis by Tian, et al.<sup>31</sup> of image-guided versus free-hand pedicle screw insertion found a total of 43 studies in the literature and reported a statistically significant lower incidence of pedicle violation with computed tomography-based navigation versus conventional insertion technique. This reduced screw deviation rate was also found in the two-dimensional and three-dimensional fluoroscopy-based navigation techniques. The authors concluded that the navigated approaches had a statistically significant higher accuracy in the placement of pedicle screws compared with conventional techniques.<sup>31</sup> In a prospective study by Laine,<sup>25</sup> 139 of 174 pedicle screws were placed with computer assistance versus free-hand technique, and reported a misplacement rate of 4.3% with computer assistance compared to a rate of 14.3% using conventional methods. In contrast, Hart, et al.<sup>32</sup> in a cadaveric study, found no significant difference in the overall thoracic pedicle screw misplacement rate between stereotactic image guidance versus manual fluoroscopic technique. Mirza, et al.<sup>26</sup> performed a similar cadaveric study evaluating the use of conventional fluoroscopy versus multiple image-guided programs. They found that accuracy was increased with the computed tomography-based image guidance, but increased procedure time by six-fold and resulted in a 20-fold increase in radiation exposure to the patient. Therefore, significantly increased operative time and radiation exposure continue to be substantial deterrents to these image-guided techniques. For these reasons, as well as several large series demonstrating minimal complications without image guidance, the free-hand technique has been espoused to be an effective and safe technique for instrumentation.<sup>15,24</sup>

During our surgical approach to thoracic spine instrumentation and fusion, we perform a wide and complete facetectomy of the inferior articular facet (IAF) bilaterally at every level. This allows for 1) complete visualization of the entire facet for morphometric confirmation of the anatomy for employment of the free-hand technique, 2) additional local bone graft material, 3) improved mobility/flexibility of the spine for deformity correction, and 4) proper preparation/decompression of the facet fusion bed. Exposing the entire SAF to its medial aspect, where the confluence of the ligamentum flavum attaches, allows the surgeon to measure accurately the width of the facet in the medial to lateral direction. From our study, if the facet measures 12 mm in width, the starting point should start 4 mm from the lateral aspect of the facet (or 2 mm lateral from the midline of the facet). The ideal cephalo-caudad starting point and sagittal inclination remains unknown and changes based on the use of a straight-forward or anatomic trajectory. We use the straight-forward trajectory, with screw placement in a position perpendicular to the superior endplate in the proximal and mid-thoracic spine and perpendicular to the lamina in the lower thoracic spine, which has been found in a cadaveric study to improve pedicle screw fixation strength.<sup>33</sup> The cephalo-caudad starting point for the straight forward trajectory as described by Kim, et al.,<sup>15</sup> changes based on the vertebral level, and is at the bisected transverse process for the first and twelfth thoracic vertebrae, trending towards a more cephalad starting point in relation to the transverse process as one proceeds towards the mid-thoracic region (T7 to T9), which has a cephalo-caudad starting point at the proximal edge of the transverse process (Table 4).

Once the starting point is determined, we use a high speed burr to create a dorsal cortical pilot hole. The pedicle probe is then inserted into the pilot hole. We prefer to use an angled pedicle probe with a tapered tip that is 2 mm in width (Figure 4). This bevel is started with the concavity oriented in the lateral direction. This should result in safe insertion of the probe. If the probe does not pass easily, the probe should

**Table 4. Cephalo-Caudad Starting Point for Straight-Forward Trajectory (Kim, et al.)<sup>15</sup>**

Thoracic level	Cephalo-caudad starting point	Medial-lateral starting point
T1	Bisected transverse process	<b>“Superior facet rule”</b> 2 mm to 3 mm lateral to midline of the superior articular facet
T2-T4	Proximal 1/3 transverse process	
T7-T9	Proximal Edge transverse process	
T10-T11	Proximal 1/3 transverse process	
T12	Bisected transverse process	

be extracted from the pilot hole, flipped to orient the bevel medially, and then the ventral lamina can be palpated medially with the use of the pedicle probe. With proper palpation of the ventral lamina, the surgeon can feel and hear the thickened cortex at the confluence of the spinal canal and medial pedicle wall, which is two to three times thicker than the lateral pedicle wall. Therefore, taking advantage of the fact that the medial cortical wall is thicker than the lateral wall, after palpating the ventral lamina, the surgeon can then “scrape” laterally along the ventral lamina and “fall” into the pedicle along the appropriate axial trajectory. The axial pedicle trajectory is nearly parallel to the sagittal plane at the twelfth thoracic vertebra and changes progressively towards a more medial trajectory with more cephalad levels. However, once the ventral lamina is palpated, and as long as the insertion of the probe stays lateral to this structure, the spinal canal should not be breached.

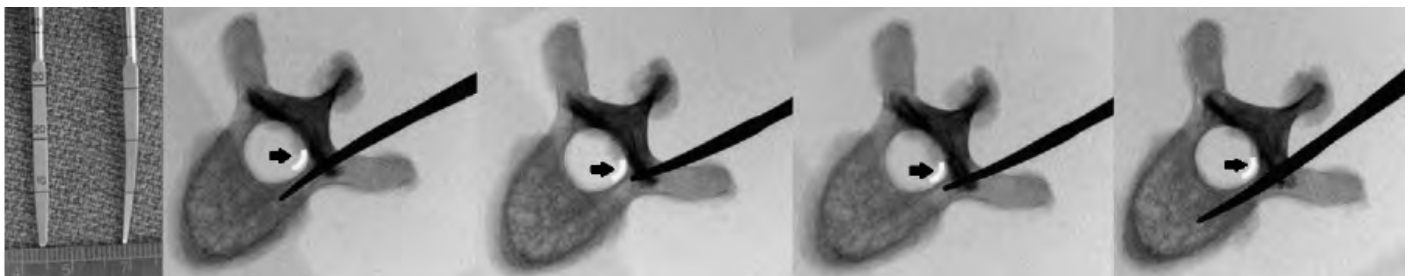
Our study has one obvious weakness, in that the cadaveric specimens evaluated did not include those with deformity. However, the majority of surgeons do not perform major deformity corrections, so the applicability of this technique and anatomic reference point applies to all surgical cases in the thoracic spine. Pedicle screw instrumentation plays a large role in deformity correction surgery, and our

findings do not specifically address whether the ventral lamina can be used as an anatomic guide in those patients with scoliosis or other pre-existing congenital deformities. While the free-hand technique has been shown to be effective even in curves > 90 degrees, it would be difficult to interpret whether the anatomic relationships associated with the ventral lamina are reproducible in the spine with deformity.<sup>22</sup> This may be of particular utility, however, as instrumenting spines with acquired or congenital deformity can be especially challenging, as pedicles may be absent, obscured, or altogether obliterated. Having stated this, the senior author (LGL) has found that the ventral lamina is a reproducible structure that aids in extremely complex deformity cases and placement of pedicle screws at the apices of curves, serving to enhance curve correction. The realization of this structure during complex deformity surgery led to initiation of the current investigation in a cadaveric model. However, the importance of the “superior facet rule” and an understanding of the ventral lamina morphometry may be particularly important for residents/fellows or an inexperienced spine surgeon, to prevent confusion and catastrophic neurologic injury, and less so for the experienced and perspicacious spine surgeon who possesses the exquisite tactile feedback to avoid intracanal violation. Therefore, given the reproducibility of the ventral lamina and “superior facet rule” at all levels in the thoracic spine, we hope our findings can be easily teachable to the training spine surgeon and will help prevent confusion and catastrophic neurologic injury compared to a more complex description, an inconsistent anatomic landmark, or a changing pedicle screw starting point based on the vertebral level.

## CONCLUSION

We have described the ventral lamina as the portion of the roof of the spinal canal that becomes confluent with the medial wall of the pedicle. Given the consistent anatomic relationship that exists between the ventral lamina and the superior articular facet at all thoracic levels, the ventral lamina can be used as a reliable guide for the medial-lateral starting point in thoracic pedicle screw placement. We recommend that the ideal medial-lateral starting point for the placement of thoracic pedicle screws start 2 mm to 3 mm lateral to the midline of the superior articular facet to avoid spinal canal violation during insertion.

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**Figure 4.** Axial radiographs of thoracic vertebrae, using an angled pedicle probe with tapered 2-mm tip, with tip pointed lateral, then re-inserted with tip pointed medial to palpate and scrape laterally along the ventral lamina (arrow) to fall into the pedicle.



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