Bone and Tendon Coverage via Dehydrated Human Amniotic/Chorionic Membrane and Split-Thickness Skin Grafting

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J Reconstr Microsurg Open 2016;1:59–62.

Extremity wounds are debilitating and complex. When critical structures are not exposed, extremity wounds can be managed with conservative measures such as local wound care or skin grafting.¹ However, exposure of critical structures such as bone or tendon precludes the use of these modalities. Skin grafts do not engraft onto bone and tendon, and with local wound care alone, bone develops osteomyelitis and tendons desiccate and rupture.² Without soft tissue reconstruction, osteomyelitis leads to amputation 50% of the time and tendon rupture results in loss of function.²

Free flap reconstruction is currently the gold standard for limb salvage. The high technical expertise required for consistent free flap limb salvage poses a barrier for patient access to care, and not all patients are suitable candidates. Additionally, microsurgical free flaps in the lower extremity have an 8% failure rate, 75% of which proceed to amputation.³ Reliable alternatives to free flap reconstruction that lead to stable wound closure are needed.

Dehydrated human amniotic/chorionic membrane (dHACM) is an FDA-approved biologic device that delivers a large number of growth factors.⁴ While dHACM is effective at closing diabetic foot ulcers and venous stasis ulcers,^{5,6} many applications of dHACM over prolonged periods are typically necessary for complete closure. The cost and time to closure makes dHACM alone an unsuitable method for limb salvage.

We developed a novel limb salvage technique using dHACM to generate granulation tissue over critical structures and then definitively closing the wound with split-thickness skin grafts (STSG). Tendon coverage requires

1 week of dHACM treatment before STSG. Bone coverage requires 2 to 3 weeks of dHACM treatment before STSG. We present our early experiences using this innovative approach.

Materials and Methods

dHACM + STSG limb salvage (►Table 1) was guided by the following parameters. Following initial debridement, wounds were assessed for size and surface area of exposed bone and/or tendon. dHACM was applied to the wounds to generate granulation tissue, and wounds were covered with a nonadherent dressing. During the ensuing week, wounds were assessed for granulation tissue. If sufficient, STSG was performed. If the granulating base was not ready, additional dHACM application was performed. Wounds were monitored carefully for infection.

Results

Between November 5, 2014, and March 30, 2015, seven patients underwent dHACM + STSG limb salvage. Demographics included 8 to 64 years of age, and two female and five male patients. Wounds included two with exposed tendons, three with exposed bone, and two with exposed bone and tendon. dHACM and STSG was successful in six of the seven patients. None developed infection during dHACM treatment, STSG, and in the postoperative phase, even in the cases where initial antibiotic treatment was inadequate due to bacterial resistance. All wounds remain stably closed.

received

June 22, 2015

accepted

November 28, 2015

published online

January 18, 2016

DOI <http://dx.doi.org/>

10.1055/s-0036-1571279.

ISSN 2377-0813.

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New York, NY 10001, USA.

Tel: +1(212) 584-4662.

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Table 1 Demographics and clinical information of our case series

Age (y)	Sex	Wound size (cm ²)	Location	Critical structure(s) exposed	No. of dHACM	Successful STSG?
8	F	16	LE	Achilles tendon	1	Yes
37	F	6	LE	Extensor tendons of foot	1	Yes
44	M	3	UE	Pectoral tendon, humerus	1	Yes
60	M	12	LE	Tibia	2	Yes
64	M	8	LE	Tibia	2	Yes
54	M	12	LE	Tibia and tibialis anterior tendon	2	Unknown
47	M	5	LE	Tibia and tibialis posterior tendon	3	Yes

Abbreviations: LE, lower extremity; UE: upper extremity.

Selected Case Reports

Case I: Foot Extensor Tendons

A 37-year-old woman whose foot was run over by a garbage truck presented with a right foot dorsum degloving injury with exposed extensor tendons and multiple closed fractures. Comorbidities included uncontrolled type 2 diabetes mellitus. The wound was treated three times with urinary bladder matrix with neither granulation tissue generation nor wound closure (► **Fig. 1**). We successfully skin grafted >75% of the wound but 6 cm² of exposed extensor tendons remained. One dHACM treatment was applied to the tendons and STSG was successful the following week. The wound remained stably closed at 2-month follow-up and the patient resumed ambulation.

Case II: Tibia with Chronic MRSA Osteomyelitis

A 64-year-old man presented with more than 20 years history of a draining midtibial wound following a motor vehicle collision. Comorbidities included diabetes, hyperlipidemia, hypertension, and chronic obstructive pulmonary disease. We performed wide soft tissue and bony debridement, resulting in 8 cm² of exposed tibia, placed dHACM, and started intravenous (IV) cefazolin (► **Fig. 2**). Three days after dHACM placement, bony cultures grew methicillin-resistant *Staphylococcus aureus* (MRSA) and switched to IV vancomycin. Notably, no clinical infection developed. Two additional dHACM treatments were applied, adequate granulation tissue developed, and STSG was successful. Of note, the superolateral corner of the wound was not treated with dHACM, resulting in persistently exposed tibia which was closed with

Fig. 1 A 37-year-old woman who was struck by a motor vehicle presented with a degloving injury of her right dorsal foot. (A) On initial referral, the patient had been treated three times with urinary bladder matrix, without coverage of her exposed extensor tendons. (B) Majority of the wound was successfully skin grafted except for central portion with exposed tendon. dHACM was applied once. (C) The wound demonstrated adequate granulation tissue 1 week later and she underwent STSG. (D) Stable closure 4 weeks following STSG.

Fig. 2 A 64-year-old man presented with chronic osteomyelitis and exposed tibia of more than 20 years following a motor vehicle collision. (A) Soft tissue and bony debridement was performed and dHACM was applied. (B) Adequate granulation tissue overlying bone after three dHACM applications. Black arrow marks bony region that was not covered by dHACM. (C) STSG was placed on granulation tissue, while a rotational flap was performed to cover the untreated bone (black arrow), demonstrating stable closure 5 weeks postoperatively, while the rotation flap required local wound care. (D) Wound appearance 2 months postSTSG.

Fig. 3 A 47-year-old man presented with Gustilo 3B injury with exposed tibia and tendon following a motorcycle versus motor vehicle collision. (A) Wound in anterior lower extremity following debridement of devitalized tissue. (B) Wound following first dHACM treatment. (C) Wound after third dHACM application; STSG was applied at this time. (D) Stable closure 1 month following successful STSG, despite extensive bony manipulation for tibia re-reduction and conversion of external fixator to ring fixator.

a local rotational flap which epidermolysed and ultimately closed with local wound care. The skin grafted portions of the wound remained stably closed at 2-month follow-up and the patient successfully resumed ambulation.

Case III: Tibia and Tibialis Posterior Tendon

A 47-year-old man presented with a Gustilo 3B injury involving 5 cm² of exposed tibia and tibialis posterior tendon following a motorcycle collision (► **Fig. 3**). Medical comorbidities included diabetes and peripheral arterial disease with single-vessel runoff on formal angiogram. Orthopedic surgery placed external fixation and negative pressure dressings, and consulted our service. We performed soft tissue debridement and three dHACM treatments, after which the wound demonstrated adequate granulation tissue STSG was successful. His closure remained stable at 1-month follow-up, despite requiring extensive subsequent bony manipulation due to loss of tibial reduction and conversion of the external fixator to a ring fixator.

Discussion

Extremity wounds with exposed critical structures, including bone and tendon, are a major burden on the American health care system with limited treatment options. Free flap reconstructions of lower extremity wounds have an increased failure rate in comparison to elective free flap procedures. Free flap reconstructions are limited due to technical expertise, access to care, and suitable patients. Although a larger sample size is needed to fully evaluate this novel treatment modality, our early experience suggests dHACM + STSG is a viable, low-cost alternative to free flap reconstruction. We have successfully applied this technique to six of seven

patients. Additional advantages to this modality include a potential decreased time of hospital stay, avoiding immobility for the patient, and need for intensive care unit monitoring, leading to decreased need of ancillary services including nursing and physical therapy. Future studies will include a randomized controlled trial, and will be aimed at optimizing patient selection, timing of treatment, and analyzing the cost utility of dHACM and STSG in comparison to free flap reconstruction in addition to understanding the biological response within the wounds.

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