

Journal of Surgical Case Reports, 2019;6, 1-4

**doi: 10.1093/jscr/rjz181** Case Report

# CASE REPORT

# Innovative treatment using tilapia skin as a xenograft for partial thickness burns after a gunpowder explosion

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# Abstract

Tilapia skin has non-infectious microbiota, high amounts of type I collagen, and similar morphological structure to human skin, so it has been suggested as a potential xenograft for the management of burn wounds. A 23-year-old male patient, with no comorbidities, arrived at our burn treatment center after a thermal injury caused by contact with flames from a gunpowder explosion. Superficial partial thickness burns were present in his right upper limb and deep partial thickness burns were present in his left upper limb. Tilapia skin was applied to the lesions, leading to complete reepithelialization within 12 and 17 days of treatment, respectively. No dressing changes were needed and no side effects were observed. Tilapia skin carries the promise of an innovative, easy-to-apply and highly available product that can become the first nationally studied animal skin registered by the National Sanitary Surveillance Agency for use in the treatment of burns.

## INTRODUCTION

According to the World Health Organization, burns are responsible for 180 000 deaths annually, the majority in low- and middleincome countries, group in which Brazil is included. Moreover, non-fatal burns account for prolonged hospitalization, disfigurement, and disability, coupled with stigma and rejection [1]. Nile Tilapia Fish Skin (NTFS) has been suggested as an option of biological material for the management of burns. The Colony Forming Units found in samples of NTFS before the process of chemical sterilization indicated the presence of normal, non-infectious microbiota [2]. NTFS also presented a large composition of type I collagen, morphology similar to human skin and high resistance and tensile extension at the break [3].

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Received: April 2, 2019. Accepted: May 17, 2019

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Despite these characteristics and their enormous significance and potential, no description is present so far in any medical literature of NTFS use in treatment of human burns.

#### CASE REPORT

A 23-year-old male patient, with no comorbidities, arrived at a burn treatment center in Fortaleza, Brazil after a thermal injury caused by contact with flames from a gunpowder explosion. Superficial partial thickness burns (SPTB) were present in right upper limb (Fig. 1) and deep partial thickness burns (DPTB) were present in left upper limb, (Fig. 2) face and anterior and posterior thorax. Involvement of 16% of total body surface area was calculated with the Lund and Browder chart. After admission as inpatient, he was resuscitated with intravenous fluids using the Parkland formula and remained hemodynamically stable. Local Institutional Review Board approval and written permission from the patient were obtained, in accordance with the Declaration of Helsinki. No conflicts of interest are present.

NTFS was subjected to a rigorous process of chemical sterilization, glycerolization and irradiation, followed by microbiological tests for bacteria and fungi, before storage in refrigerated sterile packaging. Prior to its use in the patient, the skin was washed in sterile 0.9% saline for 5 minutes, with this process being repeated three times in a row. Lima *et al.* [6], apart from describing the steps of NTFS preparation, showed



Figure 1: Superficial partial thickness burn in the right upper limb, after cleaning of the lesion.

the biomaterial did not present variations in its microscopic and tensiometric structure after chemical sterilization and irradiation and recovered its natural consistency after the rehydration process [3].

The patient was submitted to anesthesia and analgesia with 150 mg of ketamine, 10 mg of midazolam and 200 mg of tramadol. After cleaning the lesion with tap water and 2% chlorhexidine gluconate and removing necrotic and fibrinous tissue, (Fig. 3) the biomaterial was applied to the upper limbs of the patient. (Fig. 4) Coverage of at least 1 cm of healthy skin in the borders of the wound and superposition of at least 1 cm between NTFS pieces are needed, both to ensure that eventual movement in the first days of treatment will not lead to uncovering of any area of the burn. Due to the still experimental nature of the treatment with NTFS, the researchers decided to apply silver sulfadiazine cream 1%, still used as standard treatment by almost all Brazilian burn centers, to the rest of the burned areas. Finally, firm coverage of the wounds with gauze and bandage was performed.

Daily collection of blood samples was performed and no significant alterations were found. Also, vital signs and clinical status of the patient remained stable. Gauze and bandage of upper limbs were cut off every 72 hours for the first week of treatment to evaluate for NTFS adherence. The biomaterial showed good adherence to the wound bed with no need of dressing changes. (Fig. 5)





Figure 2: Deep partial thickness burn in the left upper limb, after cleaning of the lesion.

Figure 3: Process of removing necrotic and fibrinous tissue from the lesion, an essential step to allow maximal contact between NTFS and the wound bed.



Figure 4: Appearance of the left upper limb after NTFS application.



Figure 5: Appearance of the dressing on the sixth day of treatment. Good adherence of NTFS to the wound bed was detected.



Figure 6: Appearance of the right upper limb lesion after removal of NTFS, with a total of 12 days required for complete reepithelialization of the SPTB.



Figure 7: Appearance of the left upper limb lesion after removal of NTFS, with a total of 17 days required for complete reepithelialization of the DPTB.

On the 12th day of treatment, NTFS had a dried and hardened appearance and started to slough off from the patient's right upper limb. Thus, the researchers decided to remove NTFS in the area. The patient's limb was placed under a shower and the wound was soaked with water. The hydration process led to weakening, breaking and slippage of the NTFS, with exposure of the underlying healed skin. (Fig. 6) On the 17th day of treatment, a similar process was performed, allowing NTFS removal from the left upper limb. (Fig. 7) No side effects were noted.

#### DISCUSSION

Since allografts and xenografts appear to be equally effective, xenografts might be a superior choice for their increased safety and reduced price [4]. Although frog skin was previously used as a burn treatment in Brazil, it was never registered by the National Sanitary Surveillance Agency (ANVISA) [5]. Porcine skin was neither registered nationally for the use in burn wounds and has very low availability in the specialized centers. NTFS can become the first nationally studied animal skin registered by ANVISA for use in burn treatment.

When used as a xenograft for treatment of experimental burns in rats, NTFS led to improvements in the healing process and no relevant changes in hematological and biochemical parameters [6]. Tilapia collagen nano-fibers were also found to enhance skin wound healing speed in rats by promoting cell adhesion, proliferation and differentiation [7]. Another study found promising results with the application of marine collagen peptides from tilapia skin on deep partial-thickness scalds in rabbits [8]. A recent study showed tilapia collagen significantly induces epidermal growth factor and fibroblast growth factor expression, which can promote proliferation and differentiation of fibroblasts and keratinocytes [9].

Healing is expected within 2 weeks for SPTB and usually takes longer than 3 weeks for DPTB [10]. The periods of 12 and 17 days required, respectively, for complete reepithelialization of this patient's wounds in the right upper limb and the left upper limb, suggest effectiveness of NTFS as a xenograft for the treatment of human burns. Also, no dressing changes were needed in the areas covered with NTFS, due to good adherence of the biomaterial to the wound bed, suggesting its potential in reducing burn patient pain, hospital costs and overall workload of the health care team.

We recognize the limitations of analyzing these results in a single patient, but we are currently working on phase II and phase III randomized controlled trials, which will allow NTFS to firm itself as a relevant option in the therapeutic arsenal of burn wounds, producing financial and social impact for the health system.

## CONFLICTS OF INTEREST

None.

#### FUNDING

None.

#### CONFLICT OF INTEREST STATEMENT

None declared.

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