# **Case Reports**

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# Surgical approach to multiple foreign body granulomas (PMMA)

Tratamento cirúrgico seriado de múltiplos granulomas por PMMA

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# ABSTRACT

Injectable fillers have long been used for cosmetic purposes and in HIV induced lipodystrophy patients. A foreign body granuloma may be a side effect of the application of absorbable products/temporary and most often the nonabsorbable/permanent fillers. It is not possible to predict the formation of these granulomas, and it may occur regardless of the application technique. In this article we describe the surgical treatment of a patient with multiple granulomas caused by intradermal injection of PMMA in the face, with poor response to conservative treatments, and severe psychosocial impairment due to this unsightly and disfiguring complication.

Keywords: polymethyl methacrylate; granuloma; ambulatory surgical procedures

# RESUMO

Substâncias para preenchimento têm sido muito utilizadas nos últimos anos em procedimentos estéticos e para correção de lipodistrofias em pacientes HIV-positivos. O granuloma de corpo estranho pode ser secundário à aplicação de produtos absorvíveis/temporários e mais frequentemente aos preenchedores inabsorvíveis/permanentes. Não é possível prever a formação desses granulomas que podem ocorrer independentemente da técnica utilizada. Neste artigo descreve-se o tratamento cirúrgico de uma paciente com múltiplos granulomas causados por preenchimento intradérmico prévio de PMMA na face, com resposta pobre a tratamentos conservadores, e com grave comprometimento psicossocial devido a essa complicação inestética e deformante.

Palavras-chave: polimetil metacrilato; granuloma; procedimentos cirúrgicos ambulatoriais

# INTRODUCTION

In the few last years, filling substances have been widely used in aesthetic procedures and for the correction of lipodystrophy in HIV-positive patients. Products containing hyaluronic acid in different densities, bovine collagen, autologous fat, liquid silicone, poly-l-lactic acid and polymethyl methacrylate (PMMA), among others are widely used for this purpose. <sup>1</sup> According to the product's bioavailability and its chemical composition and degradation capacity, these substances can be classified as temporary or permanent, organic or inorganic and autologous or heterologous. <sup>2,3</sup> In order for a substance to be considered safe, some important aspects should be taken into account: it must be biocompatible, phagocytosis resistant, stable, non-migratory, induce the least possible inflammatory response, not be carcinogenic or teratogenic <sup>2</sup>.

Polymethyl methacrylate (PMMA) is a polymer composed of synthetic microspheres of  $30\mu$ m to  $40\mu$ m, with homogenous surface, conveyed suspended in a 1:3 solution (3.5% bovine collagen and lidocaine) or in carboxymethylcellulose colloid solution. <sup>4</sup> It should be strictly applied in the deeper layers of the dermis. <sup>1,5</sup> It initially promotes volumization, however has the capacity to stimulate neocollagenesis, with long-term effects.

The application of PMMA might be related to some adverse effects, such as nodules, inflammation, allergic reactions, dyschromias, necrosis, infection and foreign body granuloma formation. <sup>6-8.</sup> Although described in the literature as rare, these complications are often permanent and very difficult to treat.

The foreign body granuloma can be secondary to absorbable/temporary products and, more often, to nonabsorbable/permanent fillers. It is not possible to foresee the formation of these granulomas, which can occur regardless of the used technique. Granulomas often form many years after the first application.<sup>9</sup>

The present paper describes the surgical treatment of a patient bearing multiple granulomas caused by a previous PMMA based intradermal filling in the face, with a poor response to conservative treatments and severe psychosocial compromise due to this unsightly and deforming complication.

### **CASE REPORT**

A 63-year-old female patient reported having undergone facial filling 18 years before, observing the onset of palpable hardened cylindriform nodular lesions painful on palpation, located in the sites of prior application of the product. The nodules were not adhered to deep subcutaneous planes.

Ultrasonographic examination of the face showed heterogeneous elongated hypoechoic images, superficial and with defined limits, located between the skin and the subcutaneous tissue, bilaterally dispersed in the malar, periorbital, temporal and supralabial regions, in addition to the nasogenian sulcus. Dimensions varied from  $3.2 \ge 0.8 \ge 0.5$  cm in the left malar region to 3.6x 2.4 x 0.5cm in the right malar region. Doppler examination evidenced avascular images. A new ultrasound examination was performed 1 year after the first, revealing poorly-defined nodular images with the following measures: 12mm in the left nasal region; 6.6mm in the left nasogenian sulcus; 8.1mm in the superior right labial region; 11.2mm in the right nasogenian sulcus; 14.0mm in the right temporal region; and 11.0mm in the left temporal region. The examinations were performed in diverse laboratories - note that the lesions were first measured in cm and then in mm.

Magnetic resonance imaging of the face (performed with contrast) showed multiple heterogeneous nodular formations, with lobular borders and isointense signal at t1 and predominantly intense in t2 and stir, showing slight contrast uptake, with confluent appearance in the subcutaneous of the malar and mandibular regions, nasogenian sulcus, bilaterally periorbicular, possibly corresponding to foreign bodies.

The histology of one of the lesions demonstrated it was a foreign body granuloma (Figure 1).

The patient underwent multiple facial surgeries aimed at correcting the unsightly defects secondary to PMMA filling.

Due to the fact that the zygomatic region's surfaces are convex and have multiple static rhytids, a decision was made to perform the w-plasty technique, aimed at obtaining a scar with a more irregular pattern, aligned with the skin's tension lines, thus being less apparent. The w-plasty consists of triangular advancement flaps uniformly interposed, parallel to the relaxed skin tension lines (RSTL). A scalpel blade n. 11 was used, observing a minimum of 60 degrees for the triangles' angles. The closure was performed with simple stitches and nylon thread 6.0.

The nasogenian and melomentonian grooves had lesions with clinical and histological characteristics similar to those in the zygomatic region.

A choice was made to perform long incisions positioned inside the grooves so that they could be hidden in the RSTL.

A "hockey stick" incision was used in the angle of the mouth's region, allowing a more cautious divulsion of the nodules located in that area.

Surgeries with the same characteristics were performed in four surgical times, with intervals of approximately 1 month, progressing with good cicatrization, without infectious complications and satisfactory aesthetic outcome, according to the patient's report (Figures 2 and 3).



**FIGURE 1 :** Slide with 4x magnification reveals infiltrate with arrangement nodular and vacuoles, surrounded by fibrous tissue, characterizing a foreign body granuloma



FIGURE 3: Surgical times: A. Marking of the hardened area, comprising the nasogenian and melolabial sulci to the right; B. Excised granuloma bed; C. Suture along the skin's tension lines; D. Seventh postoperative day; E. Arciform marking in the nasogenian sulcus region and left labial commissure; F. Divulsion of the area to be excised; G. Removal of the nodule with hardened consistency; H. Suture along the skin's tension lines; I. Final appearance after multiple interventions for correcting unsightly defects resulting from PMMA.

# DISCUSSION

Applications of synthetic polymers in the skin generate some degree of inflammatory response, due to either the trauma caused by the application or the interaction of the receptor's tissue with the implant.

The particulate filler's biocompatibility varies according to the particles' size, shape and surface. When they are irregular, they cause greater inflammatory response, while varied sizes induce a higher degree of foreign body reaction. According to the recent literature, granuloma formation occurs in between 0.01% and 2.5% of applications.<sup>2</sup>

Three days after the PMMA injection, monocytes invade the implant, and on the sixth day they differentiate into fibroblasts. In two or three weeks the connective tissue infiltrates the material, forming clusters of the product, and the neovascularization becomes more evident. The autologous collagen fibers' density increases during the four months after the application, when the fibrosis and active vascularization.<sup>2</sup> The presence of foreign body giant cells is evident after the first week after the application, reaching its maximum number around the third week and remaining at that level for two months. The late mass development of giant cells indicates the formation of foreign body granuloma formed by multinucleated giant cells and macrophages arranged in palisade with a lymphocytic halo.<sup>1</sup>

The reason that the host reacts with a granulomatous inflammatory response in these cases is still unknown. Some authors believe that a mild chronic inflammation may occur due to improper location of the filler or even to the displacement of the material to more superficial skin layers. There are some therapeutic options for the management of this complication: intralesional corticosteroids injections, corticosteroid therapy and oral antibiotics, in addition to allopurinol. In cases that are more severe or resistant to other therapies, surgical excision is mandatory. <sup>1,2,5</sup>

Several treatments, such as oral and intralesional corticosteroids were tried in the reported case, however with absence of satisfactory response. In addition, the patient described constant pain and dissatisfaction with the lesions' aesthetic appearance. For that reason surgery was proposed to remove the greatest possible number of the most evident granulomas.

The aesthetic result of the removal of granulomas was very satisfactory, since it progressed without unsightly scars, significantly improving the nodular appearance of the analyzed regions, with a high degree of patient satisfaction. The patient continues under ambulatory follow-up, and therapy with allopurinol has been initiated to reduce small residual granulomas.<sup>4</sup>

# CONCLUSION

There is not a filling substance that can be deemed as ideal since all materials can cause early or late adverse effects. Although PMMA is relatively safe, the host's immunology is primarily responsible for the different reactions that might arise.

It is crucial that the patient be informed beforehand of the possible complications in advance of undergoing any procedure.

Currently, the trend points towards a choice for absorbable fillers, which have a lower incidence of adverse reactions and do not lead to permanent outcomes.

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