PYOGENIC GRANULOMA - REVIEW

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ABSTRACT

Pyogenic granuloma is a common non-neoplastic soft tissue growth seen on a variety sites in the body including the oral cavity. It may also occur in pregnant subjects and is referred to as a granuloma gravidarum. Although the precise etio-pathogenesis is not established, it is regarded as a reactive lesion secondary to trauma or non-specific local irritation. Hormonal stimulation during puberty, pregnancy or oral contraceptive use may also be responsible. A variety of angiogenic factors may mediate exuberant endothelial proliferation. Mostly commonly it affects adolescents and young adults with preponderance in females. The most familiar site is anterior maxillary gingivae but may also affect the mandibular gingivae, lips, tongue and buccal mucosa. Conservative surgical excision is usually curative but recurrence is not unusual. Lasers and cryotherapy may also be employed. Granulam gravidarum, however, is best left untreated until parturition.

Key words: Pyogenic granuloma, pregnancy tumor, gingiva

INTRODUCTION

Pyogenic granulomas are common non-neoplastic, localized, soft tissue growths'. The term pyogenic granuloma is a misnomer as it neither represents infection with pyogenic micro-organisms nor it is a true granuloma² However, the term has been retained in the literature because of its historic significance.

The term "Pregnancy tumor" ("Granuloma gravidarum, "Epulis gravidarum")) refers to a pyogenic granuloma occurring in the pregnancy. Such lesions usually develop during the first trimester and their incidence rises up through the seventh month of pregnancy. Nevertheless, there is no clinical or histopathologic difference between pregnancy granuloma and pyogenic granuloma that occurs in "nonpregnant" patients' Pyogenic granulomas are not exclusive to the oral cavity but are also seen on other sites in the head and neck region, trunk and extremities⁴.

This paper reviews the etio-pathogenesis, clinical features, microscopic picture, management principles and controversies in pyogenic granuloma.

ETIOLOGY AND PATHOGENESIS

The exact etiopathogenesis of oral pyogenic granuloma is not known. At present it is regarded as an endothelial proliferation of unknown cause

However, there is evidence that pyogenic granulomas result from non-specific local irritation or trauma. Possible factors include:

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- 1) Poor oral hygiene and compromised periodontal health²
- 2) Dental injuries 5
- 3) Tooth extractions particularly third molars⁶
- 4) Microtrauma due to tooth brushing'
- 5) Trauma from orthodontic appliances⁸

Hormonal influences during puberty⁹ pregnancy 10,11 and the use of oral contraceptives^{12, 13} have also been implicated.

Pyogenic granuloma has also been documented as a manifestation of graft v/s host disease after allogeneic bone marrow transplantation",

Recent research has implicated the role of angiogenic factors derived from macrophages in the rapid growth of pyogenic granulomas. Various angiogenic factors including vascular endothelial growth factor (VEGF-) ¹⁵ angiopoietin-2 (Ang-2)¹⁶ and a nitric oxide (NO) synthase protein-dependent effector mechanism" may be responsible for the rapid angiogenesis seen in pyogenic granulomas.

CLINICAL FEATURES

Oral pyogenic granulomas may develop at any age but are more frequently seen in adolescents and young adults.(Fig. **1** and 2) However, lesions may be observed in the elderly (Fig.3). There is a definitie female predilection¹⁸.¹⁹

Gingival mucosa is the most common site (75%) followed by lips, tongue and buccal mucosa. Gingival lesions have been reported to be more frequent more in the maxilla with the anterior region of both jaws being more commonly involved". (Fig.4,5, and 6). Most gingival lesions involve the labio-buccal gingivae initially but may encroach on the lingual / palatal gingivae later.

Pyogenic granuloma presents as a solitary lobulated or smooth mass, which is usually pedunculated. The colour of the lesion varies from red / purple (early) or pink (old).

Pyogenic granulomas are usually painless except for mild tenderness but tend to bleed frequently (during eating/tooth brushing)causing distress to the patients. They usually range from a few mm to several cm in size. Pyogenic granulomas may exhibit rapid growth creating a pseudo impression of a more aggressive

lesion or even malignancy.Untreated lesions may undergo fibrous maturation and the tendency to bleed may become less frequent.

HISTOPATHOLOGIC FEATURES

The lesion shows exuberant proliferation of a highly vascular elements simulating granulation tissue. The endothelial-lined vascular channels are engorged with erythrocytes. The vascular channels are composed of capillary sized vessels arranged in lobules and this characteristic arrangement is the basis to designate pyogenic granuloma as "Lobular capillary hemangioma" (LCH)²¹

The vascular channels are supported by a loose connective tissue stroma which shows a mixed inflammatory cell infiltrate with a predominance of neutrophils and histiocytes. The mitotic figures within the lesion are normal. The overlying epithelium usually shows areas of ulceration with a fibrinopurulent exudate on the surface.

Fibrous tissue becomes more prominent in untreated, mature lesions concomitant with a reduction in vascularity. This change is depicted clinically as transformation of a pyogenic granuloma into a fibroma.

TREATMENT AND PROGNOSIS

Conservative surgical excision with a scalpel is the treatment of choice and is usually curative. Alternatively electrocautery may be used to remove oral lesions in order to minimize bleeding For lesions involving the gingival / alveolar mucosa, excision down to the periosteum is advised ¹². Scaling and polishing of adjacent teeth is recommended to eliminate any local irritation. Occasionally recurrences are reported and require re-excision. Nevertheless, prognosis for intraoral pyogenic granulomas is good.

Satisfactory outcome has been reported with Laser therapy 22 ²³ and Cryosurgery²⁴ for oral pyogenic granulomas. Interestingly, multiple recurrences of intraoral pyogenic granuloma have been reportedly treated with intra-lesional corticosteroids by some 25

Lesions in pregnant subjects may require a more conservative approach and since many lesions resolve spontaneously after parturition, surgical intervention is not warranted during pregnancy unless causing significant problems. Moreover, the recurrence rate is reported to be higher for lesions removed during pregnancy probably due to the persisting hormonal influences.



Fig 1. Pyogenic Granuloma in a 10 year male



Fig 2. Granuloma Gravidarum in a 26 year female



Fig 3. Pyogenic Granuloma in a 58 year edentulous male



Fig 5 (a) Granuloma Gravidarum in a 22 year old female



Fig 4 Pyogenic Granuloma in a 30 year female



Fig. 5 (b) The excised specimen



Fig 6 (a) Pyogenic Granuloma in a 49 year male



Fig 6(c) Excised specimen



Fig 6 (b) Gingival dressing in situ after excision



Fig 6 (d) Healing 10 days post-excision

DISCUSSION

All lesions should be submitted for histopathology to rule out more serious diseases since primary as well as metastatic malignancies^{26,27,28} may present as pyogenic granulomas. Moreover, peripheral odontogenic tumors may also clinically simulate pyogenic granulomas²⁹

The use of intra-lesional corticosteroids for multiple recurrences seems questionable since substantial evidence supporting this option is lacking. Therapy with the pulsed-dye laser at vascular-specific 585 nm is very selective, usually requires no anesthesia, and produces excellent cosmetic results³⁰. Therefore, laser therapy may offer more promise in recurrent / multiple lesions but requires further clinical trials.

Pyogenic granulomas may be seen in up to 5% of pregnancies 31 . The high rate of occurrence of pyogenic granuloma in pregnant subjects warrants a thorough oral examination of all pregnant subjects since gyne-cologists and obstetricians are less familiar with this condition 32

Unfortunately, a large number of intra-oral pyogenic granulomas are treated by otolaryngologists world-wide ³³ which does not seem to be justified because of a relatively poor understanding of oral disease processes and identification of local irritants. Therefore, it is suggested that all infra-oral pyogenic granulomas may preferably be treated and followed —up in oral and maxillofacial surgery clinics.

CONCLUSIONS

Oral pyogenic granulomas are fairly common lesions and are most frequently seen on the maxillary gingivae and in pregnant subjects. Conservative surgical excision is usually curative. However, the tendency to remove these lesions during pregnancy must be resisted.

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REFERENCES

- 1 Esmeili T, Lozada-Nur F, Epstein Common benign oral soft tissue masses J Dent Clin North Am. 2005 Jan;49(1):223-40.
- 2 Neville BW, Damm DD, Allen CM, Bouqout JE "Oral and Maxillofacial Pathology". Philadelphia, WB Saunders second edition 2002.12:p448.
- 3 Ojanotko-Harri AO, Harri MP, Hurttia HM, Sewon LA: Altered tissue metabolism of progesterone in pregnancy gingivitis and granuloma. J Clin Periodontol 18: 262-266, 1991
- 4 Papageorge MB, Doku HC. An exaggerated response of intraoral pyogenic granuloma during puberty. J Clin Pediatr Dent. 1992 Spring;16(3):213-6.
- 5 Aguilo L Pyogenic granuloma subsequent to injury of a primary tooth. A case report Int J Paediatr Dent. 2002 Nov;12(6):438-41
- 6 Sapp J. Philip, Eversole R Lewis, Wysocki George P Contemporary Oral and Maxillofacial Pathology1997 9: 306
- 7 Vilmann A, Vilmann P, Vilmann H Pyogenic granuloma: evaluation of oral conditions. Br J Oral Maxillofac Surg. 1986 Oct;24(5):376-82
- 8 Kneafsey L, Hughes C. Quadhelix appliance therapy resulting in pyogenic granuloma of the tongue. Dent Update. 2002 Nov;29(9):462-3.
- 9 Papageorge MB, Doku HCAn exaggerated response of intraoral pyogenic granuloma during puberty. J Clin Pediatr Dent. 1992 Spring;16(3):213-6
- 10 Ong MA, Chai WL, Ngeow WC. Recurrent gigantic pyogenic granuloma disturbing speech and mastication: a case report and literature review. Ann Acad Med Singapore. 1998 Mar;27(2):258-61
- 11 Whitaker SB, Bouquot JE, Alimario AE, Whitaker TJ Identification and semiquantification of estrogen and progesterone receptors in pyogenic granulomas of pregnancy. J Oral Surg Oral Med Oral Pathol. 1994 Dec;78(6):755-60
- 12 Lawoyin JO, Lawoyin DO, Arowojolu MO, Lawoyin T, Akande 00. Prevalence of pregnancy related oral granuloma in a Nigerian population group and the possible role of contraceptives. Mr J Med Med Sci. 2003 Dec;32(4): 409-12.
- 13 Brooks JK. The effects of hormonal oral contraceptives on the female human periodontium and experimental animal models, a review of the literature J Baltimore Coll Dent Surg. 1980 Jun;33(2):12-6
- 14 Lee L, Miller PA, Maxymiw WG, Messner HA, Rotstein LE. Intraoral pyogenic granuloma after allogeneic bone marrow transplant. Report of three cases. Oral Surg Oral Med Oral Pathol. 1994 Nov;78(5):607-10
- 15 Freitas TM, Miguel MC, Silveira EJ, Freitas RA, Galvao HC.Assessment of angiogenic markers in oral hemangiomas and pyogenic granulomas. Exp Mol Pathol. 2005 Aug;79(1):79- 85. Epub 2005 Apr 25
- 16 Yuan K, Lin MT The roles of vascular endothelial growth factor and angiopoietin-2 in the regression of pregnancy pyogenic granuloma. Oral Dis. 2004 May;10(3):179-85

- 17 Shimizu K, Naito S, Urata Y, Sekine I, Kondo T, Katayama I.Inducible nitric oxide synthase is expressed in granuloma pyogenicum. Br J Dermatol. 1998 May;138(5): 769-73
- 18 Zain RB, Khoo SP, Yeo JF. Oral pyogenic granuloma (excluding pregnancy tumour)—a clinical analysis of 304 cases. Singapore Dent J. 1995 Jul;20(1):8-10
- 19 Booth Peter Ward Schendel A Stephen Hausamen Jarg Erich Maxillofacial Surgery 1999 Vol II 88: 1489
- 20 Al-Khateeb T, Ababneh K. Oral pyogenic granuloma in Jordanians: a retrospective analysis of 108 cases. J Oral Maxillofac Surg. 2003 Nov;61(11):1285-8
- 21 Kapadia SB, Heffner DK Pitfalls in the histopathologic diagnosis of pyogenic granuloma. Eur Arch Otorhinolaryngol. 1992;249(4):195-200
- 22 White JM, Chaudhry SI, Kudler JJ, Sekandari N, Schoelch ML, Silverman S Nd:YAG and CO2 laser therapy of oral mucosal lesions. J Clin Laser Med Surg. 1998 Dec;16(6): 299-304.
- 23 Powell JL, Bailey CL, Coopland AT, Otis CN, Frank JL, Meyer INd:YAG laser excision of a giant gingival pyogenic granuloma of pregnancy. Lasers Surg Med. 1994;14(2):178-83.
- 24 Ishida CE, Ramos-e-Silva M. Cryosurgery in oral lesions. Int J Dermatol. 1998 Apr;37(4):283-5
- 25 Parisi E, Glick P, Glick M. Recurrent intraoral pyogenic granuloma with satellitosis treated with corticosteroids. Oral Dis. 2006 Jan;12(1):70-2.
- 26 Medina BR, Barba EM, Torres AV, Trujillo SMGingival metastases as first sign of a primary uterine angiosarcoma. J Oral Maxillofac Surg. 2001 Apr;59(4):467-71
- 27 Yoshii T, Muraoka S, Sano N, Furudoi S, Komori T.Large cell carcinoma of the lung metastatic to the mandibular gingiva. J Periodontol. 2002 May;73(5):571-4
- 28 Ramon Ramirez J, Seoane J, Montero J, Esparza Gomez GC, Cerero R. Isolated gingival metastasis from hepatocellular carcinoma mimicking a pyogenic granuloma. J Clin Periodontol. 2003 Oct;30(10):926-9
- 29 Niccoli-Filho W, Gomes Mda G, Raldi FV, Seraidarian PI Peripheral ameloblastoma. J Nihon Univ Sch Dent. 1997 Mar;39(1):34-7.
- 30 Meffert JJ, Cagna DR, Meffert RM Treatment of oral granulation tissue with the flashlamp pulsed dye laser. Dermatol Surg. 1998 Aug;24(8):845-8.
- 31 Yara Teresinha Correa, Claudia Mendonca Pinto, Luiz Guilherme, Mara Lucia Senna Oliveira Marcelo Leipner de OLIVEIRA Clinical and Histological Evaluation of Granuloma Gravidarum: Case Report Braz Dent J (2000) 11(2): 135-139
- 32 Sills ES, Zegarelli DJ, Hoschander MM, Strider WE Clinical diagnosis and management of hormonally responsive oral pregnancy tumor (pyogenic granuloma). J Reprod Med. 1996 Jul;41(7):467-70
- 33 Moriconi ES, Popowich LD.Alveolar pyogenic granuloma: review and report of a case. Laryngoscope. 1984 Jun;94(6): 807-9.