BACKGROUND

Ptyalism gravidarum (PG) is a rare condition affecting pregnancies, characterized by an excessive production of saliva. Although benign and self-limited, it can cause significant discomfort during pregnancy. Very few data about PG has been published so far, which ultimately culminates in health care providers’ lack of knowledge about this disorder and its management.

OBJECTIVE AND METHODS

The aim of this work is to review all published data about ptyalism gravidarum.

A review of all articles listed in Pubmed, SCOPUS and Web of Science was performed in April 2019 using the query: “ptyalism gravidarum OR ((sialorrhea OR ptyalism OR hypersialorrhea) AND pregnancy)”. Original works, reviews, case reports and cases series concerning PG in humans were included. Articles written in languages other than English as well as articles not available online due to their age were excluded. The articles’ titles of the references of the included studies were checked to be included if related to PG.

With the initial search, a total of 110 results were obtained (36 in Pubmed, 24 in SCOPUS and 50 in Web of Science). After analysing all the articles’ titles, removing duplicates, applying the inclusion and exclusion criteria and checking references, 15 studies about sialorrhea in pregnancy were obtained. (Figure 1)

All articles were fully read and all data concerning ptyalism gravidarum was summarized in this review.

REVIEW

Definition and Epidemiology and Risk Factors

Ptyalism gravidarum is a rare condition affecting 0.05 to 0.3% of all pregnancies. It is characterized by excessive secretion of saliva. Due to its low frequency, there are no specific criteria to establish the diagnosis. It is commonly associated with nausea and vomiting during the first trimester from mild morning sickness to hyperemesis gravidarum (HG), symptoms of gastroesophageal reflux disease, as well as other eating disorders such as bulimia, anorexia nervosa or PICA Syndrome. Some studies suggest it may be more frequent in women with family history of the disorder and in psycotic patients.
Multiple pregnancies and fetal gender have been pointed as possible risk factors but the evidence is contradictory\textsuperscript{6,7}. There seems to be no association with maternal age\textsuperscript{7}.

**Clinical Presentation**

*Ptyalism gravidarum* is described as excessive salivation, up to 2L per day (versus an average of less than 1L/day on a healthy non pregnant adult). It begins during the first trimester of pregnancy, as early as the 4\textsuperscript{th} week and it may improve with time or persist until the end of pregnancy or some weeks postpartum. Patients may struggle to swallow the high amount of saliva requiring spitting cups or paper tissues to dispose redundant saliva\textsuperscript{6}. Ultimately accumulation of saliva may cause speech impairment\textsuperscript{8}.

There may be some associated symptoms such as nausea, vomiting, anorexia, bad taste, ice craving and intolerance to diverse smells or tastes. Parotid and submandibular glands may be swollen due to enlargement of the gland and local vasodilatation\textsuperscript{7}. Some patients may end up with sleep deprivation and emotional distress. Although PG symptoms usually improve during sleep time, some patients report being awaken by the excessive salivation. It’s not clear whether this entity may have any impact in weight gain during pregnancy\textsuperscript{10–12}.

**Pathophysiology**

The pathophysiology of *ptyalism gravidarum* is unknown. Some authors consider it may have a psychological origin, others believe it has an organic explanation. The improvement of symptoms during sleep is more indicative of a psychological aetiology. On the other hand, its higher prevalence during early pregnancy suggests a hormonal effect\textsuperscript{7}.

The secretion of saliva is regulated by the Autonomic Nervous System (through the glossopharyngeal and facial nerves), highly stimulated by the parasympathetic nerves which increase the amount of saliva produced\textsuperscript{9}.

As for most of the changes related to pregnancy, high hCG, progesterone and oestrogens levels may play an important role\textsuperscript{13,14}. There is some evidence that pregnancy related salivation may be mediated by Substance P like peptides segregated by the hypothalamus.
and the K1R agonist activity of the Placental Endokinin B (EKB)\textsuperscript{13}. Also, the vomiting centre and the salivary control area are anatomically close\textsuperscript{14}. All these features support a common aetiology to nausea and sialorrhea during pregnancy.

**Diagnosis**

The diagnosis of this disorder is based on its clinical presentation. Due to the fact that most of the times other conditions may also be present, when PG is suspected, physicians must always rule out simple nausea related to pregnancy, hyperemesis gravidarum, PICA and eventually more serious eating disorders such as anorexia nervosa or bulimia\textsuperscript{2,10}. No complementary tests are required. Except in the event of concomitant HG, no abnormalities are expected in biochemical or electrolyte test. Salivary gland ultrasound may reveal enlargement of these glands\textsuperscript{10}.

**Management**

Ptyalism gravidarum is a self-limited disorder associated with pregnancy. It may improve or persist during pregnancy. Usually it resolves during the second trimester or at delivery, but in rare cases it may persist as long as 1 month postpartum\textsuperscript{7}. Nevertheless, its symptoms may be quite limiting and thus pregnant women may require symptomatic treatment. It is important that every health care provider is aware of PG and tries to understand the severity of the disorder as well as its impact on patients quality of life and their ability to handle its symptoms.

There is little consensus on how to manage patients with established PG. Many therapies have been attempted to reduce symptoms, but their efficacy is questionable and highly variable.

Some conservative measures include consumption of hard candies, chewing gums, dry toasted bread or crackers, frequent drinks of still unflavoured water. Lemon drops or oral lozenges or may be used to avoid bad taste. Most of the case series report few patients submitted to hypnosis, acupuncture or other alternative treatments such as Chiropractic therapy (also known as manual therapy) resulting in no effect to complete resolution of symptoms\textsuperscript{15,16}.

Some herbal agents have also been used, including *alpinia oxyphylla*, a traditional Chinese medication used for helping digestion, antidiuresis and salivation restraint. Not only the efficacy of these treatments is inconclusive but also its safety in pregnancy is still to be determined\textsuperscript{6}.

Medication used to treat pregnancy related sickness has been widely used to treat this condition, such as the association of docylamine and pyridoxine and metoclopramide. Even though their role in improving PG symptoms is not clear, it may relief nausea which is quite commonly associated with this disorder and considered safe during pregnancy. Other pharmacologic agents have been used to reduce saliva production and/or relief symptoms such as barbiturates, anticholinergics, belladonna alkaloid, phenothiazine or scopolamine\textsuperscript{5,9}. Unfortunately the evidence of the efficacy of these treatments is sparse and most patients keep their symptoms until complete spontaneous resolution of the disorder\textsuperscript{6,17}.

**Outcomes, prognosis and recurrence**

Published data about obstetric outcomes of pregnancies complicated by PG is poor and somehow contradictory. Some authors found an increased incidence of low birth weight, probably related to low maternal weight gain and prematurity, but others have found no association with worse fetal, neonatal or maternal outcomes\textsuperscript{1,7,9}.

PG appears to be more frequent in previously affected patients, so women must be warned that they have a higher probability of suffering from this condition again in a future pregnancy\textsuperscript{7}.

**CONCLUSION**

Ptyalism gravidarum is a rare disorder associated with pregnancy that usually begins during the first trimester and persists until delivery or post-partum. Most of the few data published about the subject are case reports or case series and they consider PG not to pose any specific risks to pregnancy other than its bothersome symptoms. Its exact pathophysiology remains unknown. It’s a self-limited condition and there is no proven therapy to effectively relief symptoms or treat the disorder.

**REFERENCES**

4. Diko ML, Siewe épse Diko CN. Physico-chemistry of geophagic soils ingested to relief nausea and vomiting during preg-


ENDERECO PARA CORRESPONDÊNCIA
Manuel Gonçalves-Henriques
E-Mail: manuelghenriques@gmail.com

RECEBIDO EM: 18/06/2019
ACEITE PARA PUBLICAÇÃO: 13/10/2019