A Case Report

Effect of Periodontal Diseases on Pregnancy

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Abstract

Pregnancy is a unique physiological state that affects almost all the organs, because of changes in the hormonal milieu meant to support the pregnancy. These changes generally reversible after delivery, are sometimes not without adverse effects. Recent upsurge in the interest in periodontal disease in pregnancy is attributed to association between periodontitis and adverse pregnancy outcome. Periodontal diseases are a group of infections and conditions that cause inflammation of the gingiva and the surrounding structures, which leads to destruction of the supporting tooth structures. Periodontal infections are predominantly caused by Gram-negative bacteriae that induce local and systemic elevations of proinflammatory cytokines. Transient bacteremia that occurs due to high vascularity of the periodontal tissue may lead to direct bacterial invasion of the fetoplacental unit. The release of toxic products incites host's response and triggers an inflammatory response. As a source of subclinical and persistent infection along with the cascade of systemic inflammatory responses and immune-mediated injury, periodontitis puts the pregnancy at high risk. Evidence for and against association between oral diseases and adverse pregnancy outcome comes from crosssectional studies and a few trials. Like any other association of obstetric outcome with systemic diseases, this one is also a subject of debate. We reviewed the studies providing evidence for and against effect of periodontal diseases on pregnancy. We found that different investigators have used different parameters to define periodontal disease, hence different results. Larger randomized controlled trials with uniform definitions of disease and outcome are needed to arrive at a definite conclusion.

Keywords: Low-birth weight, Periodontal diseases, Periodontitis, Pre-eclampsia, Pregnancy, Preterm labor

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INTRODUCTION

eriodontal diseases are a group of infections that cause inflammation of the gingiva and the surrounding structures that leads to destruction of the supporting tooth structures.1 These infections are predominantly caused by Gramnega-tive, anaerobic and micro aerophilic bacteria that induce local and systemic elevations of proinflammatory prostaglandins (PGE2) and cytokines.^{2,3} Periodontitis begins with the accumulation of biofilms on the tooth surface that contain high loads of bacteriae at or below the gingival margin. 4 Toxins that are produced by these bacteria stimulate a chronic inflammatory response and lead to break-down of the periodontium, creating pockets.⁵ This further causes gingival ulcerations, alveolar bone loss and hence, toothloss. The release of toxic products from the pathogenic plaque bacteriae along with the host's response triggers an inflammatory response, putting the pregnancy at high risk.6-8 Adverse

pregnancy outcomes that have been linked to periodontal diseases include miscarriage or early pregnancy loss, low birth weight, pre-eclampsia and preterm birth (PTB). ^{6,8-11}

Literature is abounding with evidence for and against 'periodontitis is an independent risk factor for preterm birth and other adverse outcomes' hypothesis. Like any other hypothesis of association between systemic disease and obstetric outcome, this one is also a subject of debate. In this article, we have reviewed the studies addressing issues pertaining to effect of periodontal diseases on pregnancy.

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Periodontal Disease and Other Systemic Conditions

There is considerable interest in the link between oral and systemic health among dental and medical providers. Current evidence suggests that periodontal disease is associated with an increased risk for cardiovascular disease, ^{8,9} diabetes, ^{10,11} community and hospital acquired respiratory infections, ¹² and adverse pregnancy outcomes. ¹³⁻¹⁵

Individuals with periodontal disease have approximately a 1.5 – 1.9 increased odds for developing cardiovascular disease. 8,16

There appears to be a bidirectional relationship between periodontal disease and diabetes with a 2- to 3-fold increased risk for diabetes among individuals with tooth loss.

Teeth and periodontium may serve as a reservoir and may contribute to respiratory infections. Individuals with poor oral hygiene such as dental decay have a 2- to 9-fold increase odds for pneumonia.¹²

Many recent studies have reported that maternal periodontal disease may be an independent contributor to abnormal pregnancy outcomes including preterm birth, low birth weight, risk for preeclampsia, mortality, and growth restriction. However, the causality of how periodontitis influences pregnancy outcomes has not been established. 14-25

Treatment of periodontal infection may reduce the risk of other systemic conditions. In a randomized clinical trial to estimate the effect of periodontal therapy on traditional and

novel risk factors for cardiovascular disease and on markers of inflammation, D'Aiuto et al found that therapy reduced inflammatory cytokines, blood pressure, and cardiovascular risk scores.²⁶

In a small treatment trial, type 2 diabetic patients showed improved diabetic control (lower HbA1c levels) after periodontal treatment.²⁷

Several investigators have reported similar effects of oral health regimens on reduced risk for nosocomial respiratory infections. Treatment of mechanically ventilated patients with a daily oral hygiene regime consisting of an 0.12% chlorhexidine gluconate wash reduced the risk for nosocomial pneumonia. ^{28,29}

Recently, studies have been inconclusive on the effects of periodontal therapy during pregnancy for preventing adverse pregnancy outcomes. 30-32

Treatment of oral infections may represent a novel approach to improving general health.

It is estimated that over 50% of pregnant women suffer from some form of gingival disease, either gingivitis or periodontitis, with the reports of prevalence fluctuating between 30%-100% for gingivitis and 5%-20% for periodontitis.

The prevalence of periodontal diseases during

pregnancy substantiates the strategy set forth by the surgeon general, in that periodontal treatment during pregnancy may potentially improve maternal and infant health.⁵

Periodontal Disease & Its Impact on Pregnancy

Periodontal infection is one of many infections that have been associated with adverse pregnancy out comes. The hypothesis that periodontal conditions influence the outcome of a pregnancy is not a new idea. In 1931, Galloway identified that the focal infection found in teeth, tonsils, sinuses, and kidneys pose a risk to the developing fetus. His information dated back to 1916 when pregnant guinea pigs were inoculated with streptococci eluted from human stillborn fetuses. This inoculation resulted in a 100% abortion rate. To show the impact on humans, he obtained a full mouth radiographic series on 242 women presenting for prenatal care. Fifteen percent (n=57) had an apical abscess and the suggested treatment was extraction of the affected tooth.

Of those who were treated, none resulted in a miscarriage or still birth.

Galloway summarized that removal of a known focal infection, which had clearly demonstrated to be a source of danger to any pregnant woman, was more beneficial than allowing the infection to harbor throughout the pregnancy. He went on to suggest that all foci of infection should be removed early in pregnancy.³⁷

It is widely recognized that good oral health maintains the structures within the oral cavity. However, it is not universally accepted that oral health may be an independent contributor to abnormal pregnancy outcomes. Many studies have been conducted and the literature is controversial on the role periodontitis has and its influence on adverse pregnancy outcomes.

Recognition and understanding of the importance of oral health for systemic health has led to significant research into the role of maternal oral health and pregnancy outcomes. During pregnancy, changes in hormone levels promote an inflammatory response that increases the risk of developing gingivitis and periodontitis. As a result of varying hormone levels without any changes in the plaque levels, 50%-70% of all women will develop gingivitis during their pregnancy, commonly referred to as pregnancy gingivitis.

This type of gingivitis is typically seen between the second and eighth month of pregnancy.³⁸

Increased levels of the hormones progesterone and estrogen can have an effect on the small blood vessels of the gingiva, making it more permeable. ^{39,40}

This increases the mother's susceptibility to oral infections, allowing pathogenic bacteria to proliferate and contribute to inflammation in the gingiva.

This hyperinflammatory state increases the sensitivity of the gingiva to the pathogenic bacteria found in dental biofilm. Females often see these changes during other periods of their life when hormones are fluctuating, such as puberty, menstruation, pregnancy, and again at menopause.³⁹⁻⁴¹

Recent research suggests that the presence of maternal periodontitis has been associated with adverse pregnancy outcomes, such as preterm birth, 19,20,23 preeclampsia, 25 gestational diabetes, 42 delivery of a small-for-gestational-age infant, 14 and fetal loss. 43

The strength of these associations ranges from a 2-fold to 7-fold increase in risk. The increased risks suggest that periodontitis may be an independent risk factor for adverse pregnancy outcomes.

In 1996, Offenbacher et al reported a potential association between maternal periodontal infection and delivery of a preterm or low-birthweight infant.¹⁹

In a case-control study of 124 pregnant women, observations suggested that women who delivered at less than 37 weeks gestation or an infant weighed less than 2500 g had significantly worse periodontal infection than control women.

Two recent meta-analyses of the association between maternal periodontal disease and preterm birth have been published. Vergnes et al examined 17 studies and reported a pooled estimate odds ratio for preterm birth of 2.83 (95%CI:1.95-4.10,P<.0001). 5 2

Xiong et al performed a systematic review and meta-analysis of 44 studies (26 case control, 13 cohort, and 5 controlled trials) to examine the relationship between maternal periodontal disease and adverse pregnancy outcome.⁵³

The meta-analysis showed that maternal treatment of periodontal disease reduced the rate of preterm low birth weight infants as a group (pooled RR 0.53, 95% CI: 0.30-0.95.

Guidelines

Turning to the guidelines that have been produced by the Oral Health and Pregnancy Project, Prof Madianos says that oral-health professionals should be aware of six key factors:

- The importance of preserving and establishing periodontal and dental health during pregnancy, and should inform and educate their patients accordingly.
- That non-surgical periodontal therapy and dental treatment, including restorations and extractions, are safe during pregnancy and especially during the second trimester of gestation.
- That dental x-rays can be taken when needed, with the appropriate protection, and local anaesthesia can be delivered without additional risk for the foetus or the pregnant woman.
- That the use of common painkillers and systemic

- antibiotics is generally safe, although tetracyclines should be avoided.
- That, as a general rule, medication should be prescribed to the pregnant woman after communication with her obstetrician.
- That all women should receive, at the start of pregnancy, a thorough evaluation of their dental and periodontal status. When gingivitis or periodontitis are diagnosed, periodontal treatment should be provided. Periodontal therapy will improve the periodontal condition and therefore the overall health of pregnant women.

Meanwhile, obstetricians are advised to be aware of the gum changes associated with pregnancy and that periodontal disease is associated with adverse pregnancy outcomes and that they should inform pregnant women accordingly.

CONCLUSION

	dontal D		evant on Association Adverse Pregnancy C		
	Studies that for	und association or re	lationship between periodontitis and	d pregnancy outcom	es
Authors/Year Journal	Country	Study Design	Definition of Periodontal Disease	Summary	Findi
Kunnen/2007 J Clin Periodontal	Netheriands	Case Control	Healthy PD: Pocket Depths < 4mm Mild PD:1-15 tooth sites with pocket depths > 4mm and BOP Present Severe PD: > 15 tooth sites with pocket depths > 4mm and BOP present	52 woman cases: preeclampsia < 34	periodontal di prevent amon controls (82%

J Public Health Dent			defined as one or more teeth with one or more sites with probing depth > or = 4mm, loss of attachment > or = 2 mm, and bleeding on probing	of gestational diabetes (GDM) in periodontal disease	GdM twice as likely to have periodontal diseas
Xiong/2006 Am J Obstet Gynecol	US	Case Control	Periodontal Disease (PD) was defined as one or more teeth with one or more sites with probing depth > or = 4mm, loss of attachment > or = 2 mm, and bleeding on probing	NHANES III: role of periodontal disease in GDM	Women with periodont disease 3x more likely develop GDM
Cota/2006 J Periodontal	Brazil	Case Control	Periodontal Disease was 4 or more teeth with one or more sites with pocket depths > 4mm and CAL > 3mm at the same site	588 women Cases: preeclampsia	Women with periodont disease at 1.8-fold increased risk for preeclampsia
Jarjoura/2005 Am J Obstet Gynecol	US	Case Control	Presence of 5 or more sites per subject with CAL of 3mm or greater	203 women Cases: PTB/LBW	Periodontal disease associated with PTB?LBW
Geopfert/2005 Am J Obstet Gynecol	US	Case Control	Periodontal Health-no attachment loss or gingival inflammation	103 women Cases: spontaneous PTB < 32 weeks	Periodontal disease more common among vs controls

one sextant severe periodontist->5 mn of attachment
loss in any one sextant

Cankci/2004 Turkey Case Control

Aust N Z J

Obstet Gynecol

P 2 Arm that bled on probing, preclampsia and with a clinical attachment
loss 2 mn at the same site, was diagnosed as periodontal disease.

Deamoyake/1998 Thailand Case Control Periodontal health was defined using CPHTN and DMFT across Cases: LBW Case Control Periodontal State of sites with clinical attachment level 2-33 or 4 mm Periodontal disease states with Clinical Periodontal States of the Cases associated with LBW, CR 3.0 (1.39-8.33).

Santo-Pereira/2007 Brazil Cross-sectional Periodontitis was classified as 124 women Periodontal disease visual properties of the Cases associated with PTB/LBW, CR 7.5 (1.3-28.8).

Santo-Pereira/2007 Brazil Cross-sectional Early- CAL < 5 mm Amm Severe CAL > 5 mm and solar (2AL < 5 mm 5 mm 5 weeks as 124 women per motion tal disease more prevalent in women with pereira as 124 women per motion tal disease more prevalent in women with pereira as 124 women per motion tallow the per motion tallow to the per motion tallow the per mot

pocket depths > 4mm or lormore sites with BOP Moderate Severe PD: 15 or more sites with BOP Moderate Severe PD: 15 or more sites with BOP Moderate Severe PD: 15 or more sites with BOP Moderate Severe PD: 15 or more sites with BOP Mid PD: 1-15 sites with Soprecial Samples or all pathogens increase preterm birth (PTB) risk of Moderate Severe PD: 15 or more sites with BOP Moderate Severe PD

Table 1. Summary of Relevant on Association between Maternal Periodontal Disease and Adverse Pregnancy Outcomes by Study Design

Studies that found association or relationship between periodontitis and pregnancy outcome

Authors/Year Journal	Country	Study Design	Definition of Periodontal Disease	Summary	Findings
Pitiphat/2006 J Periodontal	US	Prospective	Self reported periodontitis validated by radiographs taken prior to pregnancy	101 Women	Periodontal disease may increase C-Reactive Protein levels during pregnancy
Boggess/2003 Obstet Gynecol	US	Prospective	Healthy PD: pocket depths < 4mm Mild PD: 1-15 tooth sites with pocket depths >4mm and BOP present Severe PD: > 15 tooth sites with pocket depths > 4mm and BOP present	850 Women	Periodontal disease may associated with precelampsia, OR 2.4 (1.1-5.3)
Lopez/2002 J Dent Res	Chile	Prospective Intervention Study	Presence of 4 or more teeth showing one or more sites with probing depth 4 mm or higher, and with clinical attachment loss 3 mm or higher at the same site	639Women	Periodontal disease may associated with PTB/ LBW, RR 3.5(1.5-7.9)
Jeffcoat/2001 J Am Dent Assoc	US	Prospective Observational	Periodontitis -> 3 sites with attachment loss of 3 mm or more; generalized periodontal disease 90 or more sites with attachment loss of 3 mm or more Healthy periodontium <3 sites with 3 mm of attachment loss	1313Women	Periodontal disease may associated with PTB OR 4.5(2.2-9.2)
Mitchell-Lewis/200 Eur J Oral Sci ⁵⁸	1 US	Prospective Intervention Study	Not defined	Prospective Intervention Study 164 Women	Women with PTB had higher levels of oral pathogens in mouth; PTB rate less among treated women
Lopez/2005 J Periodontal	Chile	Randomized Clinical Trial Intervention Study	Gingival Inflammation with > 25% of sites with bleeding on probing, and no sites with clinical attachment loss > 2mm	Randomized Clinical Trial of periodontal treatment among women 870 with gingivitis	Treatment significantly reduced PTB/LBW (6% among untreated vs 2% treated)
Lopez/2002 J Periodontal	Chile	Randomized Clinical Trial Intervention Study	Periodontal disease-> teeth with pocket depths > 4mm and CAL > 3mm at the same site	Randomized Clinical Trial of antepartum vs, delay periodontal treatment to reduce PTB 400 w	Periodontitis was a risk factor for PTB/LBW and theraphy reduced the rates of PTB/LBW omen

Periodontal diseases appear to be a potential risk factor for preterm birth. As well as other modifiable risk factors, these diseases must be taken in charge. Cooperation between obstetricians or general practitioners and periodontists should be developed. The promotion of the early detection and treatments of periodontal disease in young women before and during pregnancy will be beneificial especially for women at risk.

The mechanism by which these steroid hormones during pregnancy increase gingival inflammation is not known. They may have dual effects: enhancing expression of angiogenic factors and decreasing apoptosis of granuloma cells. Systemic inflammation plays a major role in the pathogenesis of preterm delivery, including pre-eclampsia, intrauterine growth restriction, and preterm delivery. Chronic infections like intrauterine infection and chorioamnionitis are linked to both preterm birth and elevated CRP levels. Furthermore, periodontal disease has been associated with increased risk of preterm low birth weight, low birth weight, and preterm birth. Periodontal intervention results in a significantly decreased incidence for preterm delivery.

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