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REVIEW ARTICLE

A REVIEW ON DIAGNOSTIC DENTAL RADIATION RISK DURING PREGNANCY: JUSTIFYING THE DILEMMA-REVIEW

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ABSTRACT

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Dental x rays, Diagnostic radiation, Pregnancy, Dental treatment protocol. Pregnancy is accompanied by a variety of physiologic, anatomic and hormonal changes that can affect the oral health. However, these patients are not medically compromised and should not be denied dental treatment simply because they are pregnant. They often pay visits to dentist for pain and infections associated with their teeth. Such patients need dental treatment and in most cases the radiograph of the involved tooth is required. It is repeatedly reported that dentists post-pone dental treatments to the period after delivery because of the insufficient knowledge of the low doses involved in diagnostic dental radiation. The delay in treatment might have adverse effects on the mother and the fetus. It is estimated that there is a one-percent increase in congenital abnormalities subsequent to an exposure of 10 rads (100 mGy) of fetal dose. Since diagnostic doses are less than 10 rads in dentistry, such abnormalities cannot be attributed to dental diagnostic doses. This article justifies with literature evidence that dental x rays are not harmful during pregnancy, provided that the dentist follows all the proper radiologic practices with respective guidelines.

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INTRODUCTION

Pregnancy causes many changes in the physiology of the female patient. These alterations are sometimes subtle but can lead to disastrous complications if proper precautions are not taken during dental treatment (Turner, 2002). Physiologically, changes occur in the cardiovascular, hematologic, respiratory, gastrointestinal, genitourinary, endocrine, and oro-facial systems (Suresh, 2004). The local physical changes occur in different parts of the body, which include the oral cavity. These collective changes may pose various challenges in providing dental care for pregnant patients and in most cases the radiograph of the involved tooth is required (Kurien, 2013)⁻ It is repeatedly reported that dentists postpone dental treatments to the period after delivery because they do not have sufficient knowledge of the low doses involved in diagnostic dental radiation^{(4).} The delay in treatment might have adverse effects on the mother and the fetus. Therefore, understanding the physiologic changes of the body and the effects of dental radiation which are used in dentistry for the pregnant women and the fetuses, is essential for the management of the pregnant mothers.

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Radiological considerations

Ionizing Radiation

The term *radiation* literally, refers to energy transmission and thus is often applied not only to x-rays, but also to microwaves, ultrasound, diathermy, and radio waves (Freeman, 1994). Of these, x-rays and gamma rays have short wavelengths with very high energy and are ionizing radiation forms. The other four energy forms have rather long wavelengths and low energy (Brent, 1999, 2009) (Freeman, 1994). The biological effects of x-rays are caused by an electro chemical reaction that can damage tissue.

According to Brent (1999, 2009), x-rays and gamma-radiation at high doses can create two types of biological effects and reproductive risks in the fetus:

Deterministic effects: These may cause congenital malformations, fetal-growth restriction, mental retardation, and abortion. Although controversial, this so-called NOAEL— No Observed Adverse Effect Level—suggests that there is a threshold dose (0.05 gray or 5 rad) (Tarsitano, 1993) below which there is no risk. It also suggests that the threshold for

gross fetal malformations is more likely to be 0.2 gray (20 rad).

Stochastic effects: These are randomly determined probabilities, which may cause genetic diseases and carcinogenesis. In this case, cancer risk is increased, and hypothetically, at even very low doses. In this sense, ionizing radiation refers to waves or particles - photons of significant energy that can change the structure of molecules such as those in DNA, or that can create free radicals or ions capable of causing tissue damage (Hall, 1991; National Research Council, 1990) (Livingston, 1998). The standard terms used are exposure (in air), dose (to tissue), and relative effective dose (to tissue). In the range of energies for diagnostic x-rays, the dose is now expressed in grays (Gy), and the relative effective dose is now expressed in sieverts (Sv) (American Academy on Pediatric Dentistry Council on Clinical Affairs Committee on the adolescent, 2008). These can be used interchangeably.

X-Ray Dosimetry

When calculating the ionizing radiation dose, such as that from x-rays, several factors to be considered include:

- Type of study,
- Type and age of equipment,
- Distance of target organ from radiation source,
- Thickness of the body part penetrated, and
- Method or technique used for the study (Wagner, 1997) (ADA, 2001).

Estimates of dose to the uterus and embryo for various frequently used radiographic examinations are summarized in Table 1.

Table 1. Comparison between various radiologic exposures
and estimated embryonic or fetal dose

Source of Radiation	Dose (mrad)
Skull	< 0.01
Cervical spine	< 0.01
Thoracic spine	0.6
Chest	0.06
Upper GI	48
KUV	263
IVP	814
Pelvis	194
Computed tomography (Gonad dose for a total scan)	0.1
Daily (cosmic) background radiation	0.01

Abbreviations: GI-gastrointestinal, KUB-kidney, ureter and bladder,

IVP-Intravenous Pyelogram.

Studies of maternal body parts farthest from the uterus, such as the head, result in a very small dose of radiation scatter to the embryo or fetus. The size of the woman, radiographic technique, and equipment performance are variable factors. Thus, data in the table serve only as guidelines. When the radiation dose for a specific individual is required, a medical physicist should be consulted. Brent (Freeman, 1994), (2009) recommends consulting the Health Physics Society website (www.hps.org) (Baykus, 2012), to view some examples of questions and answers posed by patients exposed to radiation.

Deterministic Effects of Ionizing Radiation

One potential harmful effect of radiation exposure is deterministic, which may result in abortion, growth restriction,

congenital malformations, microcephaly, or mental retardation. These deterministic effects are threshold effects, and the level below which they are induced is the NOAEL (Brent, 2009). The harmful deterministic effects of ionizing radiation have been extensively studied for cell damage with resultant embryogenesis dysfunction. These have been assessed in animal models, as well as in Japanese atomic bomb survivors and the Oxford Survey of Childhood Cancers (Sorahan, 1995) (Cetin, 2005). Additional sources have confirmed previous observations and provided more information (Groen, 2012) (Chellakooty, 2004). One is the 2003 International Commission on Radiological Protection publication, which describes biological fetal effects from prenatal irradiation and another is the Biological Effects of Ionizing Radiation-BEIR VII Phase 2 report of the National Research Council (2006) (Conrad, 2013), which discusses health risks from exposure to low levels of ionizing radiation, and new data on both epidemiological and experimental research. Mainly the ionizing radiation arises from both man-made and natural sources and at very high dose can produce damaging effects in human tissues that can be evident within few days after exposure.

Stochastic Effects of Ionizing Radiation

This refers to random, presumably unpredictable oncogenic or mutagenic effects of radiation exposure. They concern associations between fetal diagnostic radiation exposure and increased risk of childhood cancers or genetic diseases. According to Doll and Wakeford (1997) (Conrad, 2005), as well as the National Research Council (2006) BEIR VII Phase 2 report, excess cancers can result from in utero exposure to doses as low as 0.01 Sv or 1 rad. Stated another way by Hurwitz and colleagues (2006) (Conrad, 2011b), the estimated risk of childhood cancer following fetal exposure to 0.03 Gy or 3 rad doubles the background risk of 1 in 600 to that of 2 in 600 (Table 1). In one report, in utero radiation exposure was determined for 10 solid cancers in adults from age 17 to 45 years. There was a dose-response relationship as previously noted at the 0.1 Sv or 10 rem threshold. Intriguingly, nine of 10 cancers were found in females (National Research Council, 2006)⁽²⁰⁾. These likely are associated with a complex series of interactions between DNA and ionizing radiation. They also make it more problematic to predict cancer risk from low-dose radiation of less than 0.1 Sv or 10 rem. Importantly, below doses of 0.1 to 0.2 Sv, there is no convincing evidence of a carcinogenic effect (Brent, 2009; Preston, 2008; Strzelczyk, 2007).

Diagnostic Radiation

To estimate fetal risk, approximate x-ray dosimetry must be known. According to the American College of Radiology no single diagnostic procedure results in a radiation dose significant enough to threaten embryo-fetal well-being (Hall, 1991) (Farage, 2011). Radiographs in pregnancy, the AP-view chest radiograph is the most commonly used study, and fetal exposure is exceptionally small 0.0007 Gy or 0.07 mrad. With one abdominal radiograph, the embryo or fetus is directly in the x-ray beam, (Table 1). The dose is higher 0.001 Gy or 100 mrad. The standard intravenous pyelogram may exceed 0.005 Gy or 500 mrad because of several films. Most "trauma series," such as radiographs of an extremity, skull, or rib series, deliver low doses because of the fetal distance from the target area. Fetal indications for radiographic studies are limited. In some countries, x-ray pelvimetry is done for a brief presentation. Kusama et al., (Feletou, 2006), indicated that the fetus does not directly receive radiation doses during head and chest diagnostic exposures and that the absorbed dose was estimated at less than 0.01 mGy. It was concluded that in women who were unaware of their pregnancy and who had been exposed to radiation, there is no need for pregnancy termination when the exposure dose to the fetus is less than 100 mGy. Nonetheless, no radiography procedure should be carried out on pregnant women unless there is an absolute necessity. All techniques for minimizing the absorbed dose should be undertaken when such radiographs are mandated. Radiographs should be provided with well-collimated beams in precisely-protected shields. A high-kVp technique is appropriate in such cases (Gambling, 2011).

Guidelines for diagnostic imaging during pregnancy

The American College of Obstetricians and Gynecologists (2009) (Guyton, 1981), has reviewed the effects of radiographic, sonographic, and magnetic-resonance exposure during pregnancy. Its suggested guidelines are as follows.

Guidelines for Diagnostic Imaging During Pregnancy

- Women should be counseled that x-ray exposure from a single diagnostic procedure does not result in harmful fetal effects. Specifically, exposure to less than 5 rads has not been associated with an increase in fetal anomalies or pregnancy loss.
- Concern about possible effects of high-dose ionizing radiation exposure should not prevent medically indicated diagnostic x-ray procedures from being performed on a pregnant woman. During pregnancy, other imaging procedures not associated with ionizing radiation, e.g., ultrasonography, and MRI, should be considered instead of x-rays when appropriate.
- Ultrasonography and MRI are not associated with known adverse fetal effects.
- Consultation with an expert in dosimetry calculation may be helpful in calculating estimated fetal dose when multiple diagnostic x-rays are performed on a pregnant patient.
- The use of radioactive isotopes of iodine is contraindicated for therapeutic use during pregnancy.
- Radiopaque and paramagnetic contrast agents are unlikely to cause harm and may be of diagnostic benefit, but these agents should be used during pregnancy only if the potential benefit justifies the potential risk to the fetus.

Summarized from the American College of Obstetricians and Gynecologists, 2009 (Helal, 2012).

Radiographs, pregnancy and the fetus

X-rays are a type of electromagnetic radiation that have the ability to ionize the material through which it passes. Ionization of living matter results in a damage to the cells or the DNA (Henry, 2012). Depending on the amount of radiation and the stages of pregnancy, damage to the fetal cells may result in miscarriages, birth defects or mental impairment. However the dental radiation exposure of the fetus is negligible (Hibbard, 2014). The embryo and the fetus, being

much more radiosensitive than the adult counterpart, are susceptible to adverse effects which result from the radiography exposure. During the first 2 weeks after the conception, the patient may have no knowledge of being pregnant, thus making it prudent for the physician to inquire about the last menstrual period before obtaining a radiographic image. Because a general questioning does not give a definitive diagnosis about the pregnancy status, a lead shielding should be used for all the women who are in their childbearing years (Inoue, 2007). The frequency of mutations and adverse effects is directly related to the dose, and the exposure is augmented when higher than necessary radiation exposures are used to compensate for the inadequate processing quality. The exposure can also be increased, depending on the view which is taken. The radiations from the maxillary anterior views may pass through the abdominal area, with penetration from the primary beam, as well as from the scatter /radiation. Depending on the head position, a similar exposure could also occur with the posterior views. Several precautions can be taken to avoid the fetal exposure when radiographs needed (Jeffreys, 2006). Using a lead shield over the patient's abdomen, using a properly collimated beam, and using a high-speed film, can reduce the foetal exposure.

The teratogenicity of the radiation depends on the fetal age and the dose of the radiation. The greatest risk to the fetus for teratogenicity and death, is during the first 10 days after the conception (Kaňová, 2011). The most critical period of the fetal development is between 4-18 weeks after the conception. The National Commission for Radiation Protective (NCRP) (Kennedy, 2010), recommends that the cumulative fetal exposure to radiation should not exceed more than 0.20 Gy, which can cause microcephaly and mental retardation. CT is quite useful for localizing deep-seated infections and it is the modality of choice for viewing the lateral pharyngeal infections. The definition of the internal anatomy is superior to the plain film radiographs, and the bony changes are seen quite well. The CT doses are higher than those of plain radiography, but they are lower than the multiple slices for polytomography (Kenny, 2014). The CT doses depend on a variety of factors, which include the scanner type, the scanning technique, the exposure settings, the number of slices, and the slice thicknesses. The skin doses from CT can range from 0.4 to 4.7 rads, with most of the machines delivering in the 2.5 - rad range. The combined axial and coronal images require from 3.5 to 5.0 rads (Kulandavelu, 2013). However, the gonadal dose has been shown to be less for a total scan, and it ranges from 0.1 to 0.3mrads. These doses to the fetus can be kept to a minimum by carefully using the shielding devices. Moreover, if the diagnostic irradiation provides crucial information for the maintenance of the maternal and fetal viabilities, the benefits outweigh the risks of the exposure. MRI may be an alternative to CT when the fetal irradiation is considered. MRI has a greater soft tissue sensitivity and contrast as compared to CT, and thus it may help even more in the difficult cases of infections. MRI uses a magnetic field-assisted nuclear alignment in creating images. However, the risks of the fetal exposure to the strong magnetic fields are not completely known.

Oral Radiography

Oral radiography is considered safe for pregnant patients, provided protective measures such high-speed film, a lead apron and a thyroid collar are used. No increase in congenital anomalies or intrauterine growth retardation has been reported for x-ray radiation exposure during pregnancy, totaling less than 5–10 cGy and a full-mouth series of dental radiographs results in only 8×10^{-4} cGy (Leung, 2012). A bitewing and panoramic radiographic study generates about one-third the radiation exposure linked with a full-mouth series with Espeed film and a rectangular collimated beam. Patients who are concerned about radiography during pregnancy should be reassured that in all cases requiring such imaging, the dental staff will practice the ALARA (As Low As Reasonably Achievable) (Leung, 2011) principle and that only radiographs necessary for diagnosis will be obtained.

Justifying the use of *dental Xrays* in Pregnant patients

Justification 1: According to the American College of Radiology, no single diagnostic x-ray involves a radiation dose significant enough to pose a threat to the health and Normal development of the fetus (ACOG 2004) (Lindheimer, 2010).

Justification 2: More recent evidence New York State Department of Health suggests that ionizing radiation at a dose of less than 5 rad does not increase the risk of malformation, growth retardation or miscarriage (NYSDH 2006) (Lipiński, 2013).

Justification 3: NOAEL— No Observed Adverse Effect Level—suggests that there is a threshold dose (0.05 gray or 5 rad) below which there is no risk (Brent *et al* 2009) (Freeman, 1994). It also suggests that the threshold for gross fetal malformations is more likely to be 0.2 gray (20 rad).

Justification 4: According to Doll and Wakeford (1997) (Lippi, 2007), as well as the National Research Council (2006) BEIR VII Phase 2 report, excess cancers can result from in utero exposure to doses as low as 0.01 Sv or 1 rad. Stated another way by Hurwitz and colleagues (2006) (LOnberg, 2003),the estimated risk of childhood cancer following fetal exposure to 0.03 Gy or 3 rad doubles the background risk of 1 in 600 to that of 2 in 600.

Justification 5: In pregnancy, the AP-view chest radiograph is the most commonly used study, and fetal exposure is exceptionally small 0.0007 Gy or 0.07 mrad. With one abdominal radiograph, because the embryo or fetus is directly in the x-ray beam, the dose is higher 0.001 Gy or 100 mrad⁽⁵¹⁾. The National Commission for Radiation Protective (NCRP 2000)⁽⁴⁰⁾ recommends that the cumulative fetal exposure to radiation should not exceed more than 0.20 Gy, which can cause microcephaly and mental retardation. Most "trauma series," such as radiographs of an extremity, skull, or rib series, deliver low doses because of the fetal distance from the target area (Flynn *et al* 2007) (Mor, 2011).

Justification 6: However, if dental X-rays prove necessary, they should be obtained under adequate safety conditions (beam collimation, high-speed film, filter, lead protection, high kV setting or constant beams, in-use quality program), No increase in congenital anomalies or intrauterine growth retardation has been reported for x-ray radiation exposure during pregnancy, totaling less than 5–10 cGy (Muallem, 2006) and a full-mouth series of dental radiographs results in only 8×10^{-4} cGy. A bitewing and panoramic radiographic study generates about one-third the radiation exposure linked with a full-mouth series with E-speed film and a rectangular

Collimated beam and only selected peri apical or bitewing images should be contemplated in most cases.

Conclusion

Pregnancy is a unique period with many physiologic changes that leads to the formation and maturation of a new life. Every pregnant women should be encouraged to seek medical and dental care during pregnancy, as a failure in treating the developing problems affects the health of both the mother and the unborn child. The dental care professionals must gain a basic understanding of the underlying physiological changes of pregnancy, the influences which are related to the use of medications during pregnancy, radiological considerations ,and how these may interact with the delivery of dental care. This understanding aids the development of the treatment plan and the delivery of the necessary medical, nutritional and dental care, as well as it prepares the professionals for counseling their pregnant patients. Hereby we conclude that dental radiographs are considered safe to be given at any time that it is deemed necessary during pregnancy, provided that the dentist follows all the proper radiologic practices, i.e., using a radiation protective apron with a thyroid collar, using highspeed films, following the proper procedures in order to take the radiograph, and following the ALARA (As Low As Reasonably Achievable) principle (ADA 2006).

REFERENCES

- ADA Council of Dental Affairs. J Am Dent Assoc. 2001;132: 234-38.
- ADA Council of Scientific Affairs. An update on radiographic practices: information and recommendations. J Am Dent Assoc. 2001;132:234-38.
- American Academy on Pediatric Dentistry Council on Clinical Affairs Committee on the adolescent. Guidelines on oral health care for the pregnant adolescent. *Pediatr Dent*. 2008-2009; 30:102-06.
- Baykus Y, Gurates B, Aydin S, et al: Changes in serum obestatin, preptin and ghrelins in patients with gestational diabetes mellitus. *Clin Biochem* 45(3):198, 2012
- Cetin I, de Santis MS, Taricco E, et al: Maternal and fetal amino acid concentrations in normal pregnancies and in pregnancies with gestational diabetes mellitus. *Am J Obstet Gynecol* 192:610, 2005
- Chellakooty M, Vangsgaard K, Larsen T, et al: A longitudinal study of intrauterine growth and the placental growth hormone (GH)–insulin like growth factor I axis in maternal circulation. *J Clin Endocrinol Metab* 89:384, 2004
- Conrad KP, Baker VL: Corpus luteal contribution to maternal pregnancy physiology and outcomes in assisted reproductive technologies. *Am J Physiol Regul Integr Comp Physiol* 304(2):R69, 2013
- Conrad KP, Jeyabalan A, Danielson LA, et al: Role of relaxin in maternal renal vasodilation of pregnancy. *Ann N Y Acad Sci* 1041:147, 2005
- Conrad KP, Shroff SG: Effects of relaxin on arterial dilation, remodeling, and mechanical properties. *Curr Hypertens Rep* 13(6):409, 2011b
- Cooper MS: Disorders of calcium metabolism and parathyroid disease. *Best Pract Res Clin Endocrinol Metab* 25(6):975, 2011
- Dashe JS, Casey BM, Wells CE, et al: Thyroid-stimulating hormone in singleton and twin pregnancy: importance of

gestational age-specific reference ranges. *Obstet Gynecol* 106:753, 2005

- Farage MA, Maibach HI: Morphology and physiological changes of genital skin and mucosa. *Curr Probl Dermatol* 40:9, 2011
- Feldt-Rasmussen U, Mathiesen ER: Endocrine disorders in pregnancy: physiological and hormonal aspects of pregnancy. 25(6):875, 2011
- Feletou M, Vanhoutte PM: Endothelial dysfunction: a multifaceted disorder (The Wiggers Award Lecture). Am J Physiol Heart Circ Physiol 291:H985, 2006
- Freeman JP, Brand JW. Radiation doses of commonly used dental radiographic surveys. Oral Surg Oral Med Oral Pathol; 1994;77(3):285-9.
- Freemark M: Regulation of maternal metabolism by pituitary and placental hormones: roles in fetal development and metabolic programming. *Horm Res* 65:41, 2006
- Gajendra S, Kumar JV.Oral health and pregnancy: A review. N Y State Dent J. 2004;70:40-44.
- Gambling L, Lang C, McArdle HJ: Fetal regulation of iron transport during pregnancy. Am J Clin Nutr 94(6 Suppl):1903S, 2011
- Guyton AC (ed): Textbook of Medical Physiology. Philadelphia, W. B. Saunders Company, 1981, p 218
- Hauguel-de Mouzon S, Lepercq J, Catalano P: The known and unknown of leptin in pregnancy. *Am J Obstet Gynecol* 194:1537, 2006
- Hein M, Petersen Ac, Helmig RB, et al: Immunoglobulin levels and phagocytes in the cervical mucus plug at term of pregnancy. *Acta Obstet Gynecol Scand* 84:734, 2005
- Helal I, Fick-Brosnahan GM, Reed-Gitomer B, et al: Glomerular hyperfiltration: definitions, mechanisms and clinical implications. *Nat Rev Nephrol* 8(5):293, 2012
- Henry JF, Sherwin BB: Hormones and cognitive functioning during late pregnancy and postpartum: a longitudinal study. *Behav Neurosci* 126(1):73, 2012
- Henson MC, Castracane VD: Leptin in pregnancy: an update. Biol Reprod 74:218, 2006
- Hibbard JU, Shroff SG, Cunningham FG: Cardiovascular alterations in normal and preeclamptic pregnancies. In Taylor RN, Roberts JM, Cunningham FG (eds): Chesley's Hypertensive Disorders in Pregnancy, 4th ed. Amsterdam, Academic Press, 2014
- Hui C, Lili M, Libin C, et al: Changes in coagulation and hemodynamics during pregnancy: a prospective longitudinal study of 58 cases. Arch Gynecol Obstet 285(5):1231, 2012
- Inoue T, Hotta A, Awai M, et al: Loss of vision due to a physiologic pituitary enlargement during normal pregnancy. *Graefe's Arch Clin Exp Ophthalmol* 245:1049, 2007
- Jeffreys RM, Stepanchak W, Lopez B, et al: Uterine blood flow during supine rest and exercise after 28 weeks of gestation. BJOG 113:1239, 2006
- Jensen D, Wolfe LA, Slatkovska L, et al: Effects of human pregnancy on the ventilatory chemoreflex response to carbon dioxide. *Am J Physiol Regul Integr Comp Physiol* 288:R1369, 2005
- Kaňová N, Bičiková M: Hyperandrogenic states in pregnancy. Physiol Res 60(2):243, 2011
- Karakosta P, Chatzi L, Plana E, et al: Leptin levels in cord blood and anthropometric measures at birth: a systematic review and meta-analysis. *Paediatr Perinat Epidemiol* 25(2):150, 2010

- Kennedy RL, Malabu UH, Jarrod G, et al: Thyroid function and pregnancy: before, during and beyond. *J Obstet Gynaecol* 30(8):774, 2010
- Kenny L, McCrae K, Cunningham FG: Platelets, coagulation, and the liver. In Taylor RN, Roberts JM, Cunningham FG (eds): Chesley's Hypertensive Disorders in Pregnancy, 4th ed. Amsterdam, *Academic Press*, 2014
- Kulandavelu S, Whiteley KJ, Bainbridge SA, et al: Endothelial NO synthase augments fetoplacental blood flow, placental vascularization, and fetal growth in mice. *Hypertension* 61(1):259, 2013
- Kurien S, Kattimani V S, Sriram R, Sriram S K, Prabhakar Rao V K, Bhupathi A, Bodduru R, Patil N N. Management of Pregnant Patient in Dentistry. *J Int Oral Health* 2013; 5(1):88-97.
- Leung AM, Pearce EN, Braverman LD: Iodine nutrition in pregnancy and lactation. *Endocrinol Metab Clin North Am* 40(4):765, 2011
- Leung AM: Thyroid function in pregnancy. *J Trace Elem Med Biol* 26(2–3): 137, 2012
- Lindheimer MD, Kanter D: Interpreting abnormal proteinuria in pregnancy: the need for a more pathophysiological approach. *Obstet Gynecol* 115(2 Pt 1):365, 2010
- Lipiński P, Styś A, Starzyński RR: Molecular insights into the regulation of iron metabolism during the prenatal and early postnatal periods. *Cell Mol Life Sci* 70(1):23, 2013
- Lippi G, Albiero A, Montagnana M, et al: Lipid and lipoprotein profile in physiological pregnancy. *Clin Lab* 53:173, 2007
- Livingston MH, Dlllinger TM, Holder R. Considerations in the management of the pregnant patient. Spec Care Dentist; 1998;18(5):183-8.
- Löf M: Physical activity pattern and activity energy expenditure in healthy pregnant and non-pregnant Swedish women. *Eur J Clin Nutr* 65(12):1295, 2011
- LOnberg U, Damm P, Andersson AM, et al: Increase in maternal placental growth hormone during pregnancy and disappearance during parturition in normal and growth hormone-deficient pregnancies. *Am J Obstet Gynecol* 188:247, 2003
- Majed BH, Khalil RA: Molecular mechanisms regulating the vascular prostacyclin pathways and their adaptation during pregnancy and in the newborn. *Pharmacol Rev* 64(3):540, 2012
- Mandala M, Osol G: Physiological remodeling of the maternal uterine circulation during pregnancy. *Basic Clin Pharmacol* Toxicol 110(1):12, 2011
- Miehle K, Stephan H, Fasshauer M: Leptin, adiponectin and other adipokines in gestational diabetes mellitus and preeclampsia. Clin Endocrinol (Oxf) 76(1):2, 2012
- Mor G, Cardenas I, Abrahams V, et al: Inflammation and pregnancy: the role of the immune system at the implantation site. *Ann N Y Acad Sci* 1221:80, 2011
- More C, Bhattoa HP, Bettembuk P, et al: The effects of pregnancy and lactation on hormonal status and biochemical markers of bone turnover. *Eur J Obstet Gynecol Reprod Biol* 106:209, 2003
- Morikawa M, Yamada T, Turuga N, et al: Coagulationfibrinolysis is more enhanced in twin than in singleton pregnancies. *J Perinat Med* 34:392, 2006
- Muallem MM, Rubeiz NG: Physiological and biological skin changes in pregnancy. *Clin Dermatol* 24:80, 2006
- Richards AG. Dental X-ray protection. Dent Clin North Am. 1968;631-41.

- Suresh L, Radfar L. Pregnancy and lactation. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2004; ;97(6):672-82.
- Tarsitano BF, Rollings RE. The pregnant dental patient: evaluation and management. *Gen Dent* 1993;41(3):226-34.
- Taylor RN, Roberts JM, Cunningham FG (eds): Chesley's Hypertensive Disorders in Pregnancy, 4th ed. Amsterdam, Academic Press, 2014
- Teratology society public affairs committee. FDA classification of drugs for teratogenic risk. Teratology 1994; 49:446-7.
- Turner M, Aziz SR. Management of the pregnant oral and maxillofacial surgery patient. J Oral Maxillofac Surg; 2002;60:1479-88.
