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Website: www.jehp.net DOI: 10.4103/jehp.jehp 9 16

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> Received: 21-05-2016 Accepted: 16-01-2017

# **Postpartum depression risk factors:** A narrative review

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#### Abstract:

Postpartum depression is a debilitating mental disorder with a high prevalence. The aim of this study was review of the related studies. In this narrative review, we report studies that investigated risk factors of postpartum depression by searching the database, Scopus, PubMed, ScienceDirect, Uptodate, Proquest in the period 2000-2015 published articles about the factors associated with postpartum depression were assessed in Farsi and English. The search strategy included a combination of keywords include postpartum depression and risk factors or obstetrical history, social factors, or biological factors. Literature review showed that risk factors for postpartum depression in the area of economic and social factors, obstetrical history, and biological factors, lifestyle and history of mental illness detected. Data from this study can use for designing a screening tools for high-risk pregnant women and for designing a prevention programs.

Keywords:

Narrative review, postpartum depression, risk factors

#### Introduction

**)**ostpartum depression is a debilitating mental disorder with a prevalence between 5% and 60.8% worldwide.<sup>[1]</sup> The intensity of feeling inability in suffering mothers is so high that some mothers with postpartum depression comment life as the death swamp<sup>[2]</sup> while nondepressed mothers see their baby's birth as the happiest stage of their life.<sup>[3]</sup> The disease manifests as sleep disorders, mood swings, changes in appetite, fear of injury, serious concerns about the baby, much sadness and crying, sense of doubt, difficulty in concentrating, lack of interest in daily activities, thoughts of death and suicide.<sup>[4,5]</sup> Feelings of hopelessness in severe cases of illness can threaten life and lead to suicide;<sup>[6]</sup> it is a factor that causes 20% of maternal deaths in the course after giving birth.<sup>[7]</sup> In addition, issues such as fear of harming the baby (36%), weak attachment to the baby (34%) and even, in extreme cases, child suicide attempts

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have been reported.<sup>[8,9]</sup> These symptoms have serious effects on family health.<sup>[10]</sup> Therefore, susceptible people need to be identified before delivery to receive proper care measures. However, the development of screening programs as well as designing evidence-based prevention programs requires principled collection of scientific documentations. However, systematic reviews were seen in the review of some available studies that have assessed the resources in explaining the therapeutic effects of selective serotonin reuptake inhibitors on postpartum depression<sup>[11]</sup> and cognitive behavioral therapies.<sup>[12]</sup> Review studies seem to be inadequate, which evaluate the social factors besides addressing biological and psychological factors, while for achieving sufficient knowledge to design screening and preventing programs, all the factors associated with postpartum depression need be evaluated together. Thus, this study aimed to evaluate risk factors for postpartum depression during pregnancy and afterward.

How to cite this article: Ghaedrahmati M, Kazemi A, Kheirabadi G, Ebrahimi A, Bahrami M. Postpartum depression risk factors: A narrative review. J Edu Health Promot 2017;6:60.

# **Materials and Methods**

This was a review (narrative) study, in which literature in English and Farsi was evaluated using electronic search in databases of Scopus, PubMed, ScienceDirect, UpToDate, and Proquest in the time range returns between 2000 and 2015. Searching in the databases was made using key words of "postpartum depression" and "risk factors" or "predisposing factors" or "predictive factors" and "biological agents" or "social factors" or "pathophysiology" or "hormonal factors" or "lifestyle" and "pregnancy." In assessing in the PubMed database, the keywords were selected in accordance with the MeSH system. Those articles were included in the study that had done research on risk factors and predisposing factors of postpartum depression, which were of cross-sectional, cohort, case-control, interventional, and review article types. In addition, the illness diagnosis basis in these articles was the diagnosis of depression within 4 weeks after giving birth to 1 year after delivery. The articles improper regarding the adequacy of sample size, research design, and statistical methods were excluded from the study.

In the initial evaluation of the articles titles, 200 paper abstracts were extracted and evaluated by two members of the research team in terms of inclusion criteria after removal the authors' names. In case of nonagreement on the presence if inclusion criteria between the two evaluators, the articles abstract was given to the third evaluator whose opinion was determinant to include the article in the review. According to the articles arbitration, 74 papers were detected appropriate. Then, the full-texts of available articles were prepared. In case of articles with unavailable full text, correspondence was done with the authors to request them for sending the article's full text after explaining the purpose of the survey.

The articles were evaluated by three team members of the research in terms of inclusion criteria. In case of meeting the inclusion criteria, the article was reviewed and contents related to the subject were extracted. Thus, the main results of each study with the article's specifications under the relevant title were noted. After collecting, the material and content were categorized based on scientific content in their respective area subsets.

## **Results and Discussion**

Articles' assessment showed that the factors associated with postpartum depression can be classified in five domains of risk factors for psychiatric, obstetric risk factors, biological and hormonal risk factors, social risk factors, and lifestyle risk factors.

# **Psychological factors**

Previous history of depression and anxiety is among the factors that are associated with a higher risk of postpartum depression. The relationship between postpartum depression and prior onset of depression has been reported in many studies,<sup>[13,14]</sup> which has been referred to as powerful factors in postpartum depression.<sup>[15,16]</sup> The occurrence of mental health disorders such as depression during pregnancy is a powerful factor in predicting postpartum depression.<sup>[17]</sup>

There is evidence in explaining these relationships suggesting that women with a positive history of depression are more susceptible to hormonal changes.<sup>[18]</sup> In support of this finding, it has reported that a history of moderate to severe premenstrual syndrome (PMS) is a factors affecting the onset of postpartum depression.<sup>[19]</sup> In women with severe PMS, the serotonin transport system will change while the serotonin transporter polymorphism area is associated with major depression.<sup>[20]</sup> High serotonin polymorphism may lead to tryptophan depletion and induction of postpartum major depression.<sup>[21]</sup>

In addition to previous depression history, negative attitude toward the recent pregnancy, number of life events,<sup>[18]</sup> and a history of sexual abuse in the past<sup>[22]</sup> were as predisposing risk factors of postpartum depression. Furthermore, the reluctance of the baby gender<sup>[13]</sup> and having low self-esteem with the impact on parenting stress<sup>[22]</sup> are factors that contribute in the development of postpartum depression.

### **Obstetric risk factors**

Assessment the relationship between the number of delivery and postpartum depression has been associated with conflicting results. Mayberry et al. have reported postpartum depression is more prevalent in multiparous women than in nulliparous women<sup>[23]</sup> while the results of another study indicate a higher prevalence of the disease in nulliparous women.<sup>[16]</sup> Furthermore, in a study conducted by Matsin in 2013, on 86 participants within 6 weeks after delivery, it was found that having two or more children due to higher psychological burden is more likely to be associated with the occurrence of depression.<sup>[10]</sup> The discrepancies between the results of these studies suggest that the number of childbirth alone is not an independent factor for developing postpartum depression and the development of pathological conditions for the occurrence of the illness is caused by psychosocial conditions that the multiplicity of delivery creates for the women.

Risky pregnancy is also associated with an increased risk of postpartum depression. These risks include conditions that lead to performing emergency cesarean section or hospitalization during pregnancy. Postpartum complications<sup>[22,24]</sup> are also effective on the incidence of postpartum depression as much as during labor complications such as meconium passage, umbilical cord prolapse, and obstetric hemorrhages.<sup>[10]</sup> Mothers with the birth of an infant with a weight <1500 g are 4–18 times at risk for postpartum depression<sup>[25]</sup> more than others.

A mismatch between the expectations of mother and pregnancy events is as factors that affect the occurrence of depression. It has been reported that women with strong desire to have natural childbirth during the perinatal period whose delivery are done by caesarean section are more prone to risk for postpartum depression than others.<sup>[26]</sup> Spending the course of pregnancy in a natural state away from the excitements due to complications during pregnancy and preparedness for the delivery seem to be as conditions effective in the prevention of postpartum depression. Since it has been reported that the use of epidural anesthesia during childbirth, attending in childbirth preparation classes during pregnancy, and continued breastfeeding after childbirth were associated with a reduced risk of postpartum depression.<sup>[27]</sup> However, insomnia during pregnancy can lead to the risk of recurrent postpartum depression in women with a previous history of the disease.<sup>[28]</sup>

The inverse association between breastfeeding and postpartum depression shows that breastfeeding is associated with a reduction in the rate of postpartum depression. It has been reported that women exclusively breastfeed their infants in the first 3 months after childbirth show lower values of Edinburgh Postnatal Depression Scale.<sup>[29]</sup> In a study conducted by Hamdan and Tamim, it was found that breastfeeding during the first 4 months after delivery reduces the risk of postpartum depression.<sup>[30]</sup> Although no causal relationship has been established for the relationship between breastfeeding and postpartum depression, breastfeeding increases the interaction between mother and baby<sup>[31,32]</sup> and thereby may affect the health of the mother.

A relationship has been observed between low hemoglobin concentration at day 7 after delivery (<120 g/L) and postpartum depressive symptoms at day 28 after childbirth.<sup>[3]</sup> Furthermore, an effective correlation has been seen between homocysteinemia in the 1<sup>st</sup> week and 6 weeks after delivery and depression. However, there is not enough evidence in this regard that postpartum anemia can cause postpartum depression or complications of pregnancy period associated with the postpartum anemia may lead to increased risk of the disease.

#### **Biological factors**

Young age during pregnancy increases the risk of depression. The highest level of depression has been

reported in mothers aged 13–19 years<sup>[33]</sup> while the lowest rate has been seen in women with the age range of 31–35-year-old.<sup>[34]</sup> In a study conducted by Abdollahi *et al.* on 1950 women at 2–12 weeks after giving birth, it was found that increasing maternal age and maternal self-efficacy are associated with a reduced risk of postpartum depression.<sup>[35]</sup>

Studies show that glucose metabolism disorders during pregnancy are also as predisposing factors for postpartum depression so that it has been observed that women with higher blood glucose levels (mean of 120 vs. 114 mg/dl) after an hour after performing the glucose challenge test with 50 g of glucose were more at risk of postpartum depression than others.<sup>[36]</sup>

Serotonin and tryptophan levels in the blood are also known factors effective on depression. A study has shown a relationship between different serotonin transporter gene alleles and serotonin receptors with mood disorders and depression.<sup>[37]</sup> Serotonin is a monoamine neurotransmitter that is synthesized during an enzymatic route from amino acid tryptophan.<sup>[38]</sup> The amount of serotonin directly depends on the individual diet. The consumption of foods rich in protein reduces the amounts of tryptophan and serotonin in the brain while a carbohydrate snack has reverse effects.<sup>[38]</sup> In nutritional deficiencies, reduced brain tryptophan (a precursor of serotonin) up to 15% leads to increased depression scale rate of postpartum depression.<sup>[21]</sup>

Oxytocin also plays a key role in regulating emotions, social interactions, and emotional responses.<sup>[39]</sup> Higher levels of oxytocin in midpregnancy have been predictors of postpartum depression within less than the first 2 weeks after delivery.<sup>[40]</sup> Recent evidence suggests that oxytocin induces the activity of serotonin receptors<sup>[41]</sup> and reduces the response to stress. The intranasal oxytocin spraying has increased the duration of positive behaviors such as eye contact and possibility of emotions and feelings both in women and men.<sup>[42]</sup>

The role of estrogen has been also evaluated in the incidence of postpartum depression. Studies on animal models have shown that steroid and estrogen hormones are modulators of transcription from nervous neurotransmitters<sup>[43]</sup> and adjust the function of serotonin receptors.<sup>[44]</sup> This hormone causes the renewal of the generation of damaged neurons in brain and leads to the production of brain neurotransmitters.<sup>[45]</sup> In hypothalamus, estrogen also affects the neurotransmitters and regulates sleep and temperature. It has been observed that the fluctuations in this hormone or its absence is associated with depression.<sup>[45]</sup>

The role of corticotropin-releasing hormone in the regulation of steroid hormones and depression has

been studied as well. In addition to hypothalamus, this hormone is also produced during pregnancy in placenta, uterus, and ovaries and regulates the pituitary-hypothalamus-adrenal axis for production of steroid hormones.<sup>[46]</sup> After delivery and expulsion of the placenta, dramatic drop of this hormone leads to reduced production of steroid hormones such as estrogen and leads to increased susceptibility to depression in the first 12 weeks after childbirth.<sup>[47]</sup> In addition to steroid hormones, some evidence has been reported suggesting the inverse association of free thyroxine levels and total serum thyroxine concentrations with symptoms of postpartum depression.<sup>[48]</sup>

Although the relationship between thyroid dysfunction and postpartum depression has not been certainly established, the disorder may cause postpartum depression in a subgroup of women.<sup>[18]</sup> According to a report, a positive thyroid peroxidase antibody test at 32 weeks of pregnancy will increase the risk of postpartum depression as 2–3 times.<sup>[49]</sup>

In addition to the association of some endogenous hormones with postpartum depression, cytokine network and inflammatory responses have been observed to be involved in the pathophysiology of depression as well.<sup>[50]</sup> Administration of cytokines such as interferon alpha and cytokine inducers such as lipopolysaccharides and typhoid vaccines have caused behavioral changes such as mood disorders, anorexia, fatigue, sleep disorders, and other temperamental mood swings, which overlap with depression symptoms.<sup>[51]</sup> Depressed women may develop postpartum psychoneuroimmunological disorder, which is caused by inflammatory response turmoil in the normal course of labor and delivery.<sup>[52]</sup> Some evidence of changes in the regulators of T-cells has also been observed in depressed women before delivery.<sup>[53]</sup> The mechanism of explaining the changes in T-cells in depression is unknown. However, it is observed that the T-cells develop apoptosis in depressed patients. One of the possible mechanisms of explaining T-cells apoptosis in depression is the increased activity of the immune system, especially depletion of their tryptophan. Tryptophan is an essential component for the proliferation of T-cells, and in an environment free of tryptophan, the T-cells undergo apoptosis process.<sup>[51]</sup>

In depressed patients, increased apoptosis in the T-cells along with decreased response to glucocorticoids will lead to decreased available T-cells and reduced the capacity of the brain in response to immunological stimuli.<sup>[51]</sup>

#### **Social factors**

Social support refers to emotional support, financial support, intelligence support, and empathy relations.<sup>[54]</sup>The

role of social support in reducing postpartum depression has been demonstrated.<sup>[55]</sup> Reducing social support is the most important environmental factor in the onset of depression and anxiety disorders.<sup>[56]</sup> At the International Conference on Population and Development of the year, decision-making power at home and increased support of the partner have been considered as the most important solution to promote women's reproductive health.<sup>[57]</sup> The spouse sexual violence and other forms of domestic violence during pregnancy are seen as factors contributing to the incidence of postpartum depression.<sup>[58]</sup>

In addition to the women's relationship with family members and community, behaviors such as smoking during prenatal period, is of social factors associated with increased incidence of postpartum depression as 1.7 times.<sup>[59]</sup>

The simultaneous relationship between smoking and socioeconomic level and the relationship between socioeconomic level with depression complicate the association between smoking and postpartum depression. However, the physiological changes of pregnancy may seem as a stressful event for some mothers and lead to the onset of depression symptoms and start of smoking.<sup>[60]</sup>

Another social factor is employment status, especially professional careers, which have been associated with a reduced risk of postpartum depression.<sup>[61]</sup> However, education and low income are associated with the risk of postpartum depression.<sup>[36,62]</sup>

#### Lifestyle

Among the factors related to lifestyle, factors of food intake patterns, sleep status, exercise, and physical activities may affect postpartum depression. It was observed that sufficient consumption of vegetables, fruits, legumes, seafood, milk and dairy products, olive oil, and a variety of nutritious may reduce postpartum depression as 50%.<sup>[63]</sup>

Vitamin B6 is effective in the production of serotonin from tryptophan as a cofactor. Therefore, the reduction of this vitamin may be involved in the process of postpartum depression.<sup>[64]</sup> In a study, the positive relation between the level of vitamin B2 absorption at week 21 of pregnancy and postpartum depression has been reported.<sup>[37]</sup> The results of an ecological study from 23 countries found that increased seafood consumption is associated with reduced risk of postpartum depression.

The results of an ecological study on 23 countries indicated that high docosahexaenoic acid levels and increased seafood consumption have been associated with reduced risk of postpartum depression.<sup>[65,66]</sup> This compound is found in fish oil.

Among the micronutrients, reduced intake of zinc and selenium is linked with the incidence of postpartum depression.<sup>[67]</sup> It was reported in a study that zinc applies its antidepressant by influencing the serotonin reuptake.<sup>[68]</sup> Selenium deficiency is likely to affect the postpartum depression by developing thyroid dysfunction.<sup>[69]</sup> Zinc is specifically found in red meat, grains, meat, and fish.

In addition to nutritional status, sleep status is among the factors influencing the risk of depression. Evidence shows that there is a relationship between less sleep and postpartum depression.<sup>[20,70]</sup> Furthermore, an effective relationship has been observed between the rate of fatigue and depression levels in days after delivery. Periods of severe sleep deprivation have been reported in depressed women after delivery.<sup>[20]</sup> Chronic sleep deprivation affects glucose metabolism, inflammatory processes, social communications, mental health, and the quality of life.<sup>[71]</sup> In addition, acute episodes of sleep deprivation affect the immune system and increase inflammatory markers such as interleukin-6 and tumor necrosis factor while these inflammatory factors have been seen more in women with postpartum depression.

There is also some evidence to suggest that exercise and physical activity have significant benefits in reducing depression symptoms, which are comparable with medicinal benefits.<sup>[72]</sup> Moderate physical activity in the third trimester of pregnancy has lowered the postpartum depression scale at 6 weeks after the delivery.<sup>[73]</sup>

A possible mechanism is the effect of exercise on mental conditions of women by increasing the endogenous opioids and endorphins, which improve the mental health. Exercise also increases self-confidence and will eliminate negative self-assessments caused by depression. In addition, exercise will help women focusing on the environment around and solving their problems.<sup>[74]</sup>

# Conclusion

Biological factors and social factors create intertwined rings that each makes women prone to postpartum depression by affecting each other. According to the findings of this study, many biological and environmental factors, such as lifestyle-related factors, are involved in the incidence or prevention of postpartum depression through direct and indirect impact on the level of serotonin in the brain and its function. Furthermore, many environmental factors such as socioeconomic factors cause crisis conditions and postpartum depression through influencing the mental health during pregnancy. Therefore, postpartum depression prevention programs need to focus on individuals interpersonal relationships to reduce domestic violence and increase social protection in addition to modify the women's lifestyle and increase their ability to cope with the crisis conditions. Moreover, based on the results of this research, the postpartum depression predictor tools should focus on social factors and lifestyle in addition to physical health conditions of individuals.

#### Acknowledgments

This study is part of a research thesis proposal approved by the Isfahan University of Medical Sciences which was performed with financial support of the Research Council of the University.

#### Financial support and sponsorship

This study was supported by Isfahan University of Medical Sciences (Grant Number: 394313).

#### **Conflicts of interest**

There are no conflicts of interest.

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Journal of Education and Health Promotion | Volume 6 | August 2017

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