

# A Comparison of Postpartum Depression in Mothers Conceived by Assisted Reproductive Technology and Those Naturally Conceived

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## Abstract

**Background:** It is thought that mothers who conceive via assisted reproductive technology (ART) may be at greater risk of postpartum depression (PPD) because of the problems and psychological stresses associated with ART treatment. The aim of the present study is to determine the occurrence of PPD among mothers who conceive by ART in comparison with those who naturally conceive. The Edinburgh Postnatal Depression Scale (EPDS) was used to assess PPD.

**Materials and Methods:** This historical cohort study investigated 406 mothers with infants aged 3-9 months. Three hundred and eight women with natural pregnancies were selected as the control group from mothers who referred to Tehran healthcare centres for infant vaccinations. The ART group consisted of 98 women who conceived via ART at Royan Institute. Participants completed a general questionnaire that asked about education, occupation, number of children, delivery method, history of infant hospitalization, breastfeeding, mothers' and infants' ages, cause of infertility (ART group), and history of depression. A validated Persian version of the EPDS was used to measure depressive symptoms.

**Results:** The mean EPDS score in mothers who naturally conceived was  $8.38 \pm 0.35$  in comparison with mothers who conceived via ART ( $7.59 \pm 0.63$ ). The proportions of women who reported PPD were 26.0% for the control group and 20.4% for the ART group. There was no statistically significant difference in PPD between the control and ART groups ( $P=0.26$ ).

**Conclusion:** The occurrence of PPD in mothers who conceived via ART was similar to those who conceived naturally.

**Keywords:** Assisted Reproductive Technology, Edinburgh Postnatal Depression Scale, Natural Pregnancy, Postpartum Depression

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## Introduction

Pregnancy is a welcomed event, usually associated with psychological and behavioral changes, especially for women who have become pregnant by an assisted reproductive technology (ART) (1). Postpartum depression (PPD) is important because it reduces the ability of a mother to care for her infant and decreases the quality of the relationship between the mother and her infant (2). Moreover, it increases the risk of future depression for the mother, and could negatively affect the mother and child relationship (3). Kettunen et al. (4) reported that

infants having symptoms and illnesses, especially from infantile colic, were more common among depressed than nondepressed mothers.

The prevalence of PPD has been reported as 10-15% in different countries; however, in a systematic review carried out by Halbreich and Karkun, the prevalence of PPD was 0-60% in 40 countries (5). In two Iranian cities (Tabriz and Bousher) this value was determined by the Edinburgh Postnatal Depression Scale (EPDS), with estimates of 34.7 and 15.5% as reported by Sehatie Shafaei et al. (6) and Bagherzadeh et al. (7), respectively.

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In recent years, the impact of infertility on the mental health of couples has been more widely considered. Of note, infertility is a very unpleasant experience for many couples (8). Seibel and Taymor have reported that the frequency of psychological problems is 20-25% in infertile couples (9). The experience of using fertility methods is unpleasant and difficult for both couples, but it is harder for women because they must take most of the medications (10). Moreover, infertility treatments usually cause high levels of stress for couples (11). It has been shown that women who undergo ART experience anxiety and depression (12). Depression, anxiety, and health problems are common reactions at the time of medical treatments in infertile couples, with estimates of 19.1% in women and 14.6% in men. Multiple failures in these treatments can negatively affect self-esteem and increase depression symptoms in infertile women (11).

Studies have shown that mothers who conceive via ART are more emotionally vulnerable and have higher levels of distress compared to those who conceive naturally (13, 14). Fisher and colleagues have reported that those who conceive via ART show significantly more early parenting difficulties, which can negatively affect interactions between the mother and her infant. They conclude that these mothers need more support during pregnancy and after birth (15).

Based our searches, there has been no study in Iran that compared PPD in mothers who conceived via ART with those who conceived naturally. Therefore, we designed this study to use the EPDS to compare the frequency of PPD and its risk factors among these groups.

## Materials and Methods

This historical cohort study investigated 406 mothers of 3-9-month-old infants. We used convenience sampling methods for patient selection. The control group consisted of 308 mothers who had natural pregnancies and referred to the health centers affiliated with three main medical universities in Tehran, Iran for child vaccinations. The ART group consisted of mothers with 3-9-month-old infants convinced by ART and selected from the registry data bank at Royan Institute, Tehran, Iran. From these, we selected 98 mothers as the ART group.

The study was approved by the Ethics Committee of Royan Institute. All mothers signed a consent form before completing the questionnaire. The questionnaire was completed by each of the control group mothers. The ART mothers were contacted by telephone to complete the questionnaire. Questions that pertained to the mothers' and infants' ages, education, occupation, number of children, delivery method, history of infant hospitalization, breastfeeding, causes of infertility in women with infertility issues, and history of depression, along with the EPDS, were answered by each mother.

The EPDS is one of the most important screening tools used to identify PPD. It is a short, 10-item questionnaire that has a score from 0 to 30. Questions 1, 2, and 4 are scored from 0 to 3, whereas questions 3 and 5-10 are scored from 3 to 0 (16). Although it was developed for English-speaking populations, the EPDS has been validated in non-English populations. Montazeri et al. (16) validated the Persian version of the EPDS in an Iranian population and reported that the questionnaire was acceptable, reliable, and valid for this population with a Cronbach's alpha coefficient of 0.86 and test-retest reliability (interclass correlation coefficient) of 0.80. Based on the EPDS scores, we categorized the mothers into two groups according to their scores: non-depressed (score: 0-9) and depressed (score:  $\geq 10$ ). Mothers with total scores of  $\geq 10$  should be further examined for depression (17).

The sample size was calculated based on at least 4% of the clinical differences of 2.6% depressed for the ART group and 6.7% depressed for the control group in EPDS between the ART and control groups. In order to determine the sample size, the power to detect the difference and type one error were set to 0.8 and 0.05, respectively.

Data analysis was performed using the Statistical Package for the Social Sciences (SPSS) software, version 20. Results were presented as proportions (percentages) and mean  $\pm$  standard error (SE) or standard deviation (SD). One-way ANOVA, followed by Tukey and Dennett's tests for multiple comparisons, were used to compare depression scores among education levels, causes of infertility, numbers of pregnancies and breastfeedings. The chi-square test was applied to compare the numbers of depressed individuals between the control and ART groups. The t test was used to compare continuous variables between normal and depressed samples.  $P < 0.05$  were considered statistically significant.

## Results

The mean age of the mothers was  $28.87 \pm 5.18$  years (range: 17-51 years) and the infants had a mean age of  $5.37 \pm 1.30$  months (range: 3-9 months). The percentage of mothers who reported PPD were 26.0% in the control group and 20.4% in the ART group, which was not statistically significant ( $P=0.26$ ).

The mean  $\pm$  SE score for EPDS in mothers who conceived naturally was  $8.38 \pm 0.35$  and it was  $7.59 \pm 0.63$  for mothers who conceived via ART. The difference was not statistically significant ( $P=0.27$ ).

Our results showed that among mothers who conceived naturally, education level, occupation, and history of depression were significantly related to PPD. However, among mothers in the ART group, the type of delivery, history of infant's hospitalization, and history of depression had a statistical correlation with PPD (Table 1).

**Table 1:** Comparison between the studied variables and PPD scores in the control and ART groups

Variables	Naturally conceived	ART conceived
Education	P=0.01*	P=0.12
Less than diploma (n=58)	9.98 (0.88)	6.92 (1.09)
Diploma (n=131)	8.79 (0.53)	9.12 (1.04)
Higher than diploma (n=119) <sup>a</sup>	7.15 (0.52)	6.30 (1.06)
Occupation	P<0.001*	P=0.59
Employed (n=71)	5.69 (0.64)	6.95 (1.45)
Housewife (n=236)	9.14 (0.39)	7.76 (0.69)
Type of delivery	P=0.21	P=0.016*
Vaginal (n=102)	7.73 (0.59)	12.33 (3.84)
Cesarean (n=205)	8.67 (0.43)	7.44 (0.52)
History of infant hospitalization	P=0.21	P=0.016*
No (n=96)	9.04 (0.60)	10.03 (1.32)
Yes (n=210)	8.09 (0.43)	6.35 (0.61)
History of depression	P=0.015*	P=0.03*
No (n=15)	12.20 (1.63)	12.12 (1.23)
Yes (n=287)	8.21 (0.36)	7.18 (0.65)
Number of last pregnancies	P=0.99	P=0.56
One (n=277)	8.33 (0.36)	8.02 (0.77)
Two (n=18)	8.27 (1.50)	6.45 (1.08)
Three or more (n=2)	8.50 (6.50)	6.85 (2.72)
Number of children	P=0.36	P=0.13
One (n=172)	8.57 (0.47)	8.19 (0.79)
Two or more (n=131)	7.92 (0.53)	6.13 (1.0)
Baby feeding status	P=0.17	P=0.49
Breastfeeding (n=181)	8.95 (0.44)	6.93 (0.87)
Breastfeeding plus milk powder (n=91)	7.78 (0.70)	8.76 (1.37)
Milk powder (n=34)	7.29 (1.0)	7.53 (1.16)

\*; Significant at P<0.05, \*; Significant between groups with education levels of less than diploma and higher than diploma, SE; Standard error, ART; Assisted reproductive technology, and PPD; Postpartum depression. Data are presented as mean  $\pm$  SE.

Our results also showed that causes of infertility were not associated with PPD in the ART group (Table 2).

**Table 2:** Comparison of PPD scores and causes of infertility

Causes of infertility	n (%)	Mean score ( $\pm$ SE)	P value
Male factor	57 (58.2)	8.31 (0.86)	0.26
Female factor	19 (19.4)	4.73 (1.26)	
Both (male and female) factors	14 (14.3)	7.07 (1.24)	
Unexplained	8 (8.1)	10.12 (2.36)	

SE; Standard error and PPD ;Postpartum depression.

## Discussion

We evaluated the occurrence of PPD among mothers who conceived via ART in comparison with those who conceived naturally due to the impact of mothers' PPD in caring for their infants. In our study, the occurrence of PPD was 20.4% in the ART group and 26% in the control group. These results were consistent with other studies in Iran (16-18). Montazeri et al. (16) reported 22% of women with PPD at 6-8 weeks and 18% at 12-14 weeks after childbirth. Another Iranian study reported the level of PPD in women at 30% (18).

About 13-19% of mothers who have recently given birth experience depression during the postpartum period (19). These differences in the numbers of women with PPD are probably due to differences in body mass index (20), age (21), race/ethnicity (22), cultural, social and economic status, mental health perceptions, and other environmental factors (poverty, social support or perception, nutrition, stress, and biological vulnerability) (5).

PPD is a multidimensional disorder. The determination of factors (biological, psychological, and social) that predispose a mother to PPD will help identify at-risk mothers (23). One of the factors that has been suggested to increase PPD is infertility. However, in the literature, we have found three meta-analyses on PPD and none reported pregnancy via ART as a potential risk factor for PPD (24-26). In a systematic review, Ross et al. (27) showed that the risk of a higher prevalence of PPD in mothers who become pregnant via ART was very low or unchanged in comparison to those with natural pregnancies. It seemed that women who have conceived through ART usually have a more intense emotional attachment to the fetus than women with spontaneous pregnancies (28). Although the mean EPDS score in mothers who conceived naturally was slightly higher than those who conceived via ART, we found no significant difference between PPD in both groups. However, Monti et al. (29) reported that the average PPD scores in mothers who conceived via ART were higher than those who conceived naturally, but the difference between the two groups was not statistically significant by using a cut-point of more than 12. This finding was similar to those reported by Chatziandreou et al. (30) and Listijono et al. (23). The results of meta-analyses by Gressier et al. (31) showed no increased risk of significant post-partum depressive symptoms after medically assisted conception.

Many studies have reported that the risk of PPD in mother with a history of depression is more than those without any history of depression (3, 5). Silverman et al. (32) reported that women with a history of depression were 20 times more likely to have PPD. The results of another study showed that the risk of PPD in women who had depression before delivery was 6 times higher than those without depression (33). The results of our study showed that the history of depression in mothers in both groups (control and ART) had a significant relationship with PPD, which was consistent with the results reported by Silverman et al. (32), Sadr et al. (34), and Davé et al. (35).

Our study showed a significant relationship between PPD and type of delivery in mothers who conceived by ART, which did not agree with the results reported by Sadr et al. (34). These findings supported the results of a study by Rahmani et al. (36). Kettunen et al. (4) reported a relationship between complicated delivery and PPD, especially with pain during delivery.

Vigod et al. (37) reported in their systematic review that mothers who gave birth to an infant of very pre-term or very low birth weight (LBW) had higher levels

of depression throughout the first postpartum year. The results of our study showed a significant relationship between infant hospitalization and PPD in mothers who conceived by ART. Mothers in the ART group appeared to worry more about their infants than the control group mothers because of the difficulties with conceiving (infertility, cost of infertility treatment).

The results of our study showed no significant relationship between occupational and educational status in both groups. Two previous studies have shown that working mothers probably have a protective factor for PPD (38, 39). Lewis et al. (40) observed that employed women reported less symptoms of depression than stay-at-home mothers, regardless of their weekly working hours. Sadr et al. (34) showed that there was no significant relationship between PPD and occupational and educational status.

Limitations of this study included the relatively low sample size in the ART group and limited access to this patient population since some of these women lived in other cities and only referred to Royan Institute for infertility treatment before pregnancy. In addition, we did not have consents from all of the women who became pregnant after ART at Royan Institute and could not enroll them in this study. Further studies with larger sample sizes are recommended.

## Conclusion

This study reveals that the occurrence of PPD in mothers who conceived naturally is similar to those who conceived via ART. Our study has also provided evidence that levels of education, occupation, type of delivery, history of infant hospitalization, and history of depression are risk factors for PPD in mothers. These factors, rather than conception via ART, should be given further prominence in interventions to prevent PPD in women.

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## Authors' Contributions

F.M., E.A.: Conception and design of study, data collection, and drafting the manuscript. Z.E., Data collection. Sh.S., M.Ch., F.M., E.A., Z.E.: Analysis and/or interpretation of data. All authors read and approved the final version of the manuscript.

## References:

1. Su TJ, Tzeng YL, Kuo PC. The anxiety of Taiwanese women with or without continuity treatment after previous in vitro fertilization failure. *J Clin Nurs*. 2011; 20 (15-16): 2217-2223.
2. O'Higgins M, Roberts IS, Glover V, Taylor A. Mother-child bonding at 1 year: associations with symptoms of postnatal depression and bonding in the first few weeks. *Arch Womens Ment Health*. 2013;

- 16(5): 381-389.
3. Paulson JF, Bazemore SD. Prenatal and postpartum depression in fathers and its association with maternal depression: a meta-analysis. *JAMA*. 2010; 303(19): 1961-1969.
4. Kettunen P, Koistinen E, Hintikka J. The connections of pregnancy-, delivery-, and infant-related risk factors and negative life events on postpartum depression and their role in first and recurrent depression. *Depress Res Treat*. 2016; 2016: 2514317.
5. Halbreich U, Karkun S. Cross-cultural and social diversity of prevalence of postpartum depression and depressive symptoms. *J Affect Disord*. 2006; 91(2-3): 97-111.
6. Sehhatie Shafaei F, Ranjbar koochaksarie F, Ghojzadeh M, Mohamadrezaei Z. Study of relationship between some predisposing factors and postpartum depression. *J Ardabil Univ Med Sci*. 2008; 8(1): 54-61.
7. Bagherzadeh R, Zahmatkeshan N, Motamed N, Khoramroudi R, Ganjoo M. Prevalence of maternal blues, postpartum depression and their correlation with premenstrual syndrome in Women Referred to Health Centers Affiliated to Bushehr University of Medical Sciences. *Iranian Journal of Obstetrics, Gynecology and Infertility*. 2009; 12(3): 9-15.
8. Guerra D, Liobra A, Veiga A, Barri PN. Psychiatric morbidity in couples attending a fertility service. *Hum Reprod*. 1998; 13(6): 1733-1736.
9. Seibel MM, Taymor ML. Emotional aspects of infertility. *Fertil Steril*. 1982; 37(2): 137-145.
10. Domar AD, Smith K, Conboy L, Iannone M, Alper M. A prospective investigation into the reasons why insured United States patients drop out of in vitro fertilization treatment. *Fertil Steril*. 2010; 94(4): 1457-1459.
11. Repokari L, Punamäki RL, Poikkeus P, Vilksa S, Unkila-Kallio L, Sinkkonen J, et al. The impact of successful assisted reproduction treatment on female and male mental health during transition to parenthood: a prospective controlled study. *Hum Reprod*. 2005; 20(11): 3238-3247.
12. Kahyaoglu Sut H, Balkanli Kaplan P. Quality of life in women with infertility via the fertiQOL and the hospital anxiety and depression scales. *Nurs Health Sci*. 2015; 17(1): 84-89.
13. Cecotti M. Medically assisted procreation. Psychological aspects of infertility, parenthood and parentage. Rome: Armando Editore; 2004.
14. Hart VA. Infertility and the role of psychotherapy. *Issues Ment Health Nurs*. 2002; 23(1): 31-41.
15. Fisher JR, Hammarberg K, Baker HW. Assisted conception is a risk factor for postnatal mood disturbance and early parenting difficulties. *Fertil Steril*. 2005; 84(2): 426-430.
16. Montazeri A, Torkan B, Omidvari S. The Edinburgh Postnatal Depression Scale (EPDS): translation and validation study of the Iranian version. *BMC Psychiatry*. 2007; 4: 7: 11.
17. Cox JL, Holden JM, Sagovsky R. Detection of postnatal depression: development of the 10-item Edinburgh Postnatal Depression Scale. *Br J Psychiatry*. 1987; 150: 782-786.
18. Mazaheri MA, Rabiei L, Masoudi R, Hamidzadeh S, Nooshabadi MR, Najimi A. Understanding the factors affecting the postpartum depression in the mothers of Isfahan city. *J Educ Health Promot*. 2014; 3: 65.
19. Le Strat Y, Dubertret C, Le Foll B. Prevalence and correlates of major depressive episode in pregnant and postpartum women in the United States. *J Affect Disord*. 2011; 135(1-3): 128-138.
20. LaCoursiere DY, Barrett-Connor E, O'Hara MW, Hutton A, Varner MW. The association between prepregnancy obesity and screening positive for postpartum depression. *BJOG*. 2010; 117(8): 1011-1018.
21. Matsumoto K, Tsuchiya KJ, Itoh H, Kanayama N, Suda S, Matsuzaki H, et al. Age-specific 3-month cumulative incidence of postpartum depression: the Hamamatsu Birth Cohort (HBC) study. *J Affect Disord*. 2011; 133(3): 607-610.
22. Liu CH, Tronick E. Rates and predictors of postpartum depression by race and ethnicity: results from the 2004 to 2007 New York City PRAMS survey (Pregnancy Risk Assessment Monitoring System). *Matern Child Health J*. 2013; 17(9): 1599-1610.
23. Listijono DR, Mooney S, Chapman M. A comparative analysis of postpartum maternal mental health in women following spontaneous or ART conception. *J Psychosom Obstet Gynaecol*. 2014; 35(2): 51-54.
24. O'Hara MW, Swain AM. Rates and risk of postpartum depression-a meta-analysis. *Int Rev Psychiatry*. 1996; 8(1): 37-54.
25. Beck CT. Predictors of postpartum depression: an update. *Nurs Res*. 2001; 50(5): 275-285.

26. Robertson E, Grace S, Wallington T, Stewart DE. Antenatal risk factors for postpartum depression: a synthesis of recent literature. *Gen Hosp Psychiatry*. 2004; 26(4): 289-295.
27. Ross LE, McQueen K, Vigod S, Dennis CL. Risk for postpartum depression associated with assisted reproductive technologies and multiple births: a systematic review. *Hum Reprod Update*. 2011; 17(1): 96-106.
28. McMahon CA, Boivin J, Gibson FL, Hammarberg K, Wynter K, Saunders D, Fisher J. Age at first birth, mode of conception and psychological wellbeing in pregnancy: findings from the parental age and transition to parenthood Australia (PATPA) study. *Hum Reprod*. 2011; 26(6): 1389-1398.
29. Monti F, Agostini F, Fagandini P, La Sala GB, Blickstein I. Depressive symptoms during late pregnancy and early parenthood following assisted reproductive technology. *Fertil Steril*. 2009; 91(3): 851-857.
30. Chatziandreu M, Madianos MG, Farsaliotis VC. Psychological and personality factors and in vitro fertilization treatment in women. *Eur J Psychiatry*. 2003; 17(4): 223-231.
31. Gressier F, Letranchant A, Cazas O, Sutter-Dallay AL, Falissard B, Hardy P. Post-partum depressive symptoms and medically assisted conception: a systematic review and meta-analysis. *Hum Reprod*. 2015; 30(11): 2575-2586.
32. Silverman ME, Reichenberg A, Savitz DA, Cnattingius S, Lichtenstein P, Hultman CM et al. The risk factors for postpartum depression: A population-based study. *Depress Anxiety*. 2017; 34(2): 178-187.
33. Ongerli L, Otieno PA, Mbui J, Juma E, Mathai M. Antepartum risk factors for postpartum depression: a follow up study among urban women living in Nairobi, Kenya. *J Preg Child Health*. 2016; 3: 5.
34. Sadr SS, Dowlatian M, Behboudi Moghadam Z. Prevalence of postpartum depression and factors affecting it in Tehran. *Journal of Medical Council of IRI*. 2004; 22(3): 189-193.
35. Davé S, Petersen I, Sherr L, Nazareth I. Incidence of maternal and paternal depression in primary care: a cohort study using a primary care database. *Arch Pediatr Adolesc Med*. 2010; 164(11): 1038-1044.
36. Rahmani F, Seyedfatemi N, Asadollahi M, Seyedrasooli A. Predisposing factors of postpartum depression. *Iran Journal of Nursing*. 2011; 24 (72): 78-77.
37. Vigod SN, Villegas L, Dennis CL, Ross LE. Prevalence and risk factors for postpartum depression among women with preterm and low-birth-weight infants: a systematic review. *BJOG*. 2010; 117(5): 540-550.
38. Gjerdingen D, McGovern P, Attanasio L, Johnson PJ, Kozhimannil KB. Maternal depressive symptoms, employment, and social support. *J Am Board Fam Med*. 2014; 27(1): 87-96.
39. Goyal D, Gay C, Lee KA. How much does low socioeconomic status increase the risk of prenatal and postpartum depressive symptoms in first-time mothers? *Womens Health Issues*. 2010; 20(2): 96-104.
40. Lewis BA, Billing L, Schuver K, Gjerdingen D, Avery M, Marcus BH. The relationship between employment status and depression symptomatology among women at risk for postpartum depression. *Womens Health (Lond)*. 2017; 13(1): 3-9.