## Association of Postpartum Depression with Maternal Serum Magnesium Levels, Infant Growth, and Neurodevelopmental Indices

#### **Abstract**

Background: Postpartum depression (PPD) can exert both short-term and long-term effects on a child's health. Offspring born to mothers who suffer from PPD face an elevated susceptibility to encountering psychological disturbances and developmental delays. Moreover, there has been conjecture surrounding a plausible connection between maternal magnesium (Mg) levels and psychiatric manifestations. This study aims to investigate the relationship between maternal Mg levels and PPD and the correlation between PPD and an infant's growth and neurodevelopment at 6 and 12 months, Methods: This longitudinal study is a sub-study derived from the "PERSIAN Birth Cohort Study," encompassing 224 mother-infant pairs randomly enlisted during 2019-2020 in Isfahan. Maternal serum magnesium (Mg) levels were measured at 38 weeks of gestation. PPD was evaluated employing the Edinburgh Postpartum Depression Scale (EPDS) four weeks postpartum. Measurements of birth size were undertaken, adhering to standardized protocols at birth, 6 months, and 12 months. Anthropometric parameters and the Persian version of the validated Ages and Stages Questionnaires (ASQ) were employed to assess infant neurodevelopmental status at 6 and 12 months. Results: Overall, 22.3% of mothers grappled with PPD. The mean (standard deviation) maternal magnesium levels (Mg) were 1.95 ± 0.23 mg/dL. No statistically significant association was detected between maternal serum magnesium (Mg) levels and the incidence of PPD. Correspondingly, no significant association emerged between PPD and indices of growth. However, a noteworthy distinction materialized in the communication scores of offspring born to depressed and non-depressed mothers following adjustments for confounding variables at 12 months ( $\beta = 1.81$ ; 95% confidence interval: 0.32-3.30). Furthermore, a substantial regression in communication skills became apparent between 6 and 12 months. Conclusions: This study failed to establish a significant association between maternal serum magnesium (Mg) levels and PPD. Nevertheless, research lends credence to an inverse correlation between maternal depression and subsequent behavioral difficulties in offspring, such as communication skills. Thus, the imperative nature of screening for PPD should be underscored to facilitate its early detection and intervention, thereby enhancing infant well-being.

Keywords: Child development, infant, magnesium, postpartum depression

#### Introduction

Postpartum depression (PPD), a subtype of major depressive disorder, manifests as persistent and severe low mood occurring within four weeks after childbirth and potentially extending up to 30 weeks postpartum. **Symptoms** encompass crying episodes, insomnia, despondency, fatigue, anxiety, impaired concentration, diminished interest or pleasure in activities, reduced energy levels, and feelings of worthlessness or guilt.[1] PPD's prevalence is estimated to range between 10% and 15% in North America,[2] 27% in the Middle East, and 23.7%-34.8% in  $Iran.^{[3,4]}$ 

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The role of trace elements in the PPD. pathophysiology of including magnesium (Mg),has garnered attention.<sup>[5]</sup> Magnesium, the body's fourth most abundant cation (Mg<sup>2+</sup>), is pivotal in activating enzyme systems. Magnesium deficiency is associated with psychiatric symptoms such as depression, headaches, tremors, anxiety, and mood swings.[6,7] The substantial transfer of magnesium (Mg<sup>2+</sup>) from the mother to the fetus can lead to issues in PPD.[8] Magnesium (Mg) supplementation has shown efficacy in treating depression.[6,7]

Conversely, untreated maternal depression can have short-term and long-term repercussions on the child's well-being.

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Neonates born to depressed mothers are less likely to be breastfed and are more prone to immune problems, sleep disturbances, impaired fetal and postnatal development, and increased hospitalizations.<sup>[1,9,10]</sup>

The first year of a child's life is a critical period for growth and development, influenced by the mother's emotional state. There have been conflicting findings regarding the association between PPD and anthropometric measures. Avan *et al.*'s<sup>[11]</sup> study highlighted an association between PPD and stunting in Africa or a higher height-for-age z-score and adiposity in depressed women in the United States.<sup>[12]</sup> Conversely, Grote *et al.*'s<sup>[13]</sup> study concluded that postnatal depression does not significantly affect offspring growth in European countries.

Moreover, several studies have demonstrated a substantial impact of maternal PPD on infant development at various stages. A systematic review indicated that a mother's mood in the first year of a child's life can negatively influence four domains of the child's development: behavior, psychomotor skills, cognition, and social–emotional development.<sup>[14]</sup> Consequently, children born to mothers with PPD are at risk of cognitive and language developmental delays. [9,15] Some studies have also highlighted developmental delays across multiple domains in children due to maternal mood, encompassing problem-solving, communication, personal–social skills, and motor skills.<sup>[14,16]</sup>

PPD represents a relatively prevalent mood disorder following childbirth, and many sufferers refrain from seeking treatment from mental health professionals. Furthermore, while magnesium (Mg) has been posited as a contributing factor, the available data remain inconclusive. In light of PPD's adverse impact on maternal responsibilities and interaction with infants, the current study aimed to investigate two hypotheses: first, whether there exists an association between maternal magnesium (Mg<sup>2+</sup>) levels and PPD, and second, whether there is a relationship between PPD and infants' growth and neurodevelopmental indicators.

#### **Methods**

#### Study population

This longitudinal study was conducted as a sub-study within the framework of the Prospective Epidemiological Research Studies in Iran (PERSIAN Birth Cohort) at the Isfahan Center, known as the "Isfahan Birth Cohort" (IBC). Detailed information regarding the study's design, participants, and data collection methods has been previously published.<sup>[17]</sup>

Sample size estimation for this study was determined using the correlation coefficient formula, with a significance level of 5% and a statistical power of 80%. [18] Initially, the minimum required sample size for this study was calculated to be 199 subjects. However, considering the potential for

a reduction in sample size, it was conservatively increased to 219. Ultimately, data from 224 mother-infant pairs were included and subjected to analysis.

All participants were selected through a random process utilizing random table numbers. These participants were provided with comprehensive information about the study's objectives, protocols, and subsequent follow-up stages. To participate, they willingly provided their informed consent by signing consent forms and completing relevant questionnaires. Furthermore, participants were informed about the necessity of blood sampling during the later stages of pregnancy, with an assurance that their personal information would be kept strictly confidential.

#### **Inclusion criteria**

The pregnant women enrolled in this study were Iranian residents of Isfahan for at least one year, with no prior history of infertility and the intention to deliver their babies in hospitals within Isfahan City.

#### **Exclusion criteria**

Women who had relocated to another city or country or those who had consumed alcohol or cigarettes during pregnancy were ineligible for participation in this study. Furthermore, the study excluded mothers experiencing preterm labor, preeclampsia, a history of unfortunate events such as divorce or the death of a relative, those taking psychotropic medication, or individuals undergoing other stressful life experiences.

#### **Ethical consideration**

The study protocol, which adhered to the criteria of the Helsinki Declaration, was submitted to the Ethics Committee of the Isfahan University of Medical Science and received approval under the ethical code IR.MUI.MED. REC.1398.331. Participation in this study was entirely voluntary, and participants retained the right to withdraw from the study at any point.

#### Sample collection

Maternal blood samples, totaling 2 ml and collected in Heparinized tubes, were obtained at the 38th week of pregnancy. Following blood preparation, serums were stored in a refrigerator at temperatures ranging from 2 to 8°C using a xylidyl blue calorimeter. Serum magnesium levels were subsequently determined at the laboratory of Amin Hospital, affiliated with Isfahan University of Medical Science, utilizing the absorbance photometry method with the Mindray BS-600 Analyzer.

#### Birth size measurements

A proficient interviewer gathered general, medical, and socioeconomic data, including education, income, parental occupations, and smoking habits. Subsequently, a skilled midwife employed standardized techniques to obtain neonatal anthropometric measurements, encompassing

weight (in grams), head circumference, and height (in centimeters), conducted at least two hours post-birth. Birth weight was assessed using a precision instrument that could measure within 10 grams (Seca mobile digital baby scale 334; UK). Birth length was determined using a stadiometer, with the neonate lying supine and their knees extended while the soles of their feet were securely positioned against the measurement board. Head circumference was measured at the maximal occipitofrontal circumference (OFC) using a standard measuring tape that was accurate to 1 mm. The mid-upper arm circumference, defined as the circumference of the upper left arm, was measured at the midpoint between the shoulder and elbow at 6 and 12 months of age using colored plastic tape.<sup>[19]</sup>

#### **Data collection**

Gestational age, neonatal sex, gravidity (the number of pregnancies a woman has had), parity (the number of viable children a woman has given birth to), maternal weight (in kg), maternal height (in cm), maternal supplementation during pregnancy (yes or no), delivery method (vaginal, cesarean, or other), maternal employment status (housewife or employed), breastfeeding status, and socioeconomic status (SES) categorized as low, moderate, or high were meticulously recorded.

Furthermore, the initial demographic information questionnaire administered to mothers upon entry into the birth cohort was utilized to gather essential details and evaluate SES.

Gestational age was determined based on the mother's regular menstrual cycle or the first-trimester fetal ultrasound (performed at or before 16 weeks of gestation), employing crown-rump length as the measurement criterion.

Additionally, maternal body mass index (BMI) at enrollment was calculated by dividing body weight (in kg) by the square of height (in m²), maternal education level (<12 years, 12 years [high school diploma], or >12 years), monthly household income (in Iranian Rials [IRR]), and exposure to second-hand smoke during pregnancy (yes or no) were documented.

#### **Outcome measures**

#### Study questionnaire

The EPDS comprises ten questions, each offering four response options to assess maternal moods and feelings. It is administered during the fourth week following delivery. Every question is allocated a score ranging from 0 to 3, resulting in a total score ranging from 0 to 30. A score of 12 or higher indicates the presence of PPD. [20,21] Existing evidence supports the sufficient validity and reliability of the EPDS in the Iranian population for diagnosing PPD. [22] Information concerning maternal dietary supplements, including magnesium, was documented during the third trimester of pregnancy.

The Persian version of the validated Questionnaires of Ages and Stages (ASQ) is employed to evaluate the neurodevelopmental status of the child at 6 and 12 months.<sup>[23]</sup> It screens for developmental delays in children utilizing 20 age-specific developmental questionnaires, commencing at 2 months and concluding at 60 months. These questionnaires encompass five domains: Fine Motor, Gross Motor, Communication, Problem-Solving, and Personal-Social. Each domain consists of six questions, with response options of yes (10 points), sometimes (5 points), or not yet (0 points). This study utilized the questionnaires designed for the 6- and 12-month age groups.<sup>[24]</sup> The test's validity ranges from 76% to 88%, rendering it a reliable tool with a Cronbach's alpha coefficient of 0.86 and a reliability coefficient of 0.93 for Iranian children. Its specificity and sensitivity were 95% and 90%, respectively, for the high-risk group and 100% and 75% for the community group. [25]

#### Statistical analysis

Descriptive statistics, encompassing frequency (%) and mean (SD), were utilized to illustrate participants' categorical and continuous data. The two-independent sample t-test (in instances of non-normal data, the Mann-Whitney test was employed), along with the Chi-square test, was employed to ascertain disparities in maternal and/or infant demographic characteristics between infants born to postpartum depressed and non-depressed mothers. Logistic regression analysis was employed to explore the association between maternal serum magnesium levels and the risk of PPD. The model underwent adjustments for maternal weight, height, delivery method, maternal age at delivery, SES gravidity, parity, household income, and infant gender. The researchers utilized repeated measures analysis of variance to assess changes in anthropometric measurements (z-scores) and skill measurements from 6 to 12 months among infants born to postpartum depressed and non-depressed mothers.

Additionally, correlation and multivariate regression analyses were conducted to evaluate the association between PPD and anthropometric (z-scores) and neurodevelopmental skills at 12 months while controlling for 6-month scores and other covariates. These covariates encompassed delivery type, maternal age, SES, parity, maternal education, maternal employment, breastfeeding at 12 months, exclusive breastfeeding, supplementation, infant gender, and birth weight. Statistical significance was established for all analyses at a *P* value less than 0.05. All statistical analyses were executed using IBM SPSS Statistics V. 22.0 (SPSS Inc., Chicago, IL).

#### **Results**

The frequency of PPD among the participants stood at 22.3%. The mean age of the participants in our study was  $28.23 \pm 4.34$  years. The mean maternal serum magnesium

level was  $1.95 \pm 0.23$  mg/dl. No significant association was observed between maternal serum magnesium levels and the risk of PPD (odds ratio [OR]: 0.29, 95% confidence interval [CI]: 0.04 to 1.77). This statistical model was adjusted for maternal weight, height, delivery type, maternal age at delivery, SES, number of previous deliveries, previous children, maternal education, and infant gender. Comprehensive characteristics of mothers

and their children, stratified by PPD, can be found in Table 1.

There were no significant differences detected in infants' weight-for-height, weight-for-age, height-for-age, and mid-upper arm circumference-for-age at 6 and 12 months, based on PPD status (P > 0.05) [Table 2]. Furthermore, there was no significant depression-by-time effect (interaction effect) on infants' weight-for-height, weight-for-age, height-for-age,

Table 1: Characteristics of mothers and children according to postpartum depression						
Demographic characteristics	Non-depressed (n=174)	Depressed (n=50)	P			
Maternal magnesium level (mg/dl), mean (SD)	1.97 (0.25)	1.91 (0.18)	0.64			
Maternal weight (kg), mean (SD)	65.67 (11.57)	64.81 (11.10)	0.63			
Maternal height (cm), mean (SD)	161.38 (5.20)	161.80 (5.49)	0.09			
Gravidity (number of pregnancy)						
The first labor	78 (75.7%)	25 (24.3%)	0.64			
Second	66 (81.5%)	15 (18.5%)				
Third or more	30 (76.9%)	9 (23.1%)				
Socioeconomic status (SES)						
Low	58 (68.2%)	27 (31.8%)	0.02*			
Moderate	93 (83.8%)	18 (16.2%)				
High	22 (84.6%)	4 (15.4%)				
Parity (number of labor with live child)						
None	78 (75.7%)	26 (24.3%)	0.91			
One	70 (78.7%)	15 (21.3%)				
Two or more	24 (80.0%)	9 (20.0%)				
Type of delivery						
Vaginal	84 (80.0%)	21 (20.0%)	0.76			
Cesarean	82 (76.6%)	25 (23.4%)				
Other	8 (72.7%)	3 (27.3%)				
Maternal employment						
Housewife	162 (76.8%)	49 (23.2%)	-			
Employee	11 (100%)	-				
Maternal education						
Illiterate/under 12th education	21 (56.8%)	16 (43.2%)	0.004*			
Twelfth education	74 (81.3%)	17 (18.7%)				
Academic	78 (82.1%)	17 (17.9%)				
Children						
Newborn weight (gr)	3161.79 (379.29)	3161.84 (399.81)	0.99			
Newborn height (cm)	50.20 (2.44)	50.53 (2.59)	0.41			
Newborn head circumference (cm)	34.60 (1.33)	34.69 (1.21)	0.67			
Breastfeeding at 12 months						
No	14 (82.4%)	3 (17.6%)	0.6			
Yes	158 (77.8%)	45 (22.2%)				
Exclusive breastfeeding						
No	74 (76.3%)	23 (23.7%)	0.55			
Yes	98 (79.9%)	25 (20.3%)				
Supplementation						
No	54 (83.1%)	11 (16.9%)	0.22			
Yes	120 (75.5%)	39 (24.5%)				

<sup>\*</sup>P<0.05 is considered statistically significant. Data are shown as mean (SE) or frequency (%). Results were reported from an two-independent sample t-test and Chi-square test

or mid-upper arm circumference-for-age (P > 0.05) [Table 3]. The overall change in anthropometric measurements from 6 to 12 months did not exhibit a significant difference between depressed and non-depressed mothers (P > 0.05) [Table 2]. However, a significant depression-by-time effect (interaction effect) was observed in communication skills, indicating a poorer progression in communication skills for children of depressed mothers compared to non-depressed mothers (P = 0.02) [Table 3].

Furthermore, as depicted in Table 4, a multivariate linear model demonstrated a significant association between PPD status and 12-month communication skills in both models a ( $\beta$ : 1.89; 95% CI: 0.44-3.34) and model b ( $\beta$ : 1.81; 95% CI: 0.32-3.30) [Table 4].

#### **Discussion**

The present study assessed the associations between maternal magnesium levels and PPD. Additionally, the association between PPD and a child's growth and neurodevelopment at 6 and 12 months was investigated. Findings indicate no significant differences in the growth of children born to depressed mothers compared to non-depressed mothers at birth, 6 months, or 12 months. However, a significant association between PPD status and communication skills at 12 months was observed after controlling for confounding variables. The response rate for this study was 93.4%.

The rate of PPD in our study was lower than in previous studies conducted in our country<sup>[4,26]</sup> but higher than reported in North America.<sup>[2]</sup> This disparity may be attributed to various influential factors on depression before

and after childbirth, such as obstetric, pediatric, personality, socioeconomic, and cultural factors. Additionally, using different validated questionnaires and cut-off values for assessing depression has led to inconsistent results. [26,27] Furthermore, in alignment with a study by Smorti *et al.*, [28] it was observed that the likelihood of experiencing depression was higher among mothers with lower SES and less education than their higher socioeconomic and more educated counterparts. Poverty and socioeconomic hardships are recognized as risk factors for depression. [29]

Our study's mean maternal magnesium levels were comparable to those observed in pregnant women deficiency.[30,31] without magnesium However, significant association was found between maternal serum magnesium levels and PPD. This finding aligns with a randomized controlled trial by Edalati Fard, which showed that the prescription of magnesium supplements did not reduce postpartum depressive symptoms.[32] The association between decreased magnesium levels and depression in various groups has not been substantiated.[33] Several factors, including hormonal, physiological, and psychological changes, appear to play a more significant role in the etiology of depression, particularly during the prenatal period, than maternal serum magnesium levels. [27,34]

Our study revealed no differences in child growth between infants born to depressed and non-depressed mothers at birth, 6 months, or 12 months. Although the weight-for-age z-score at 12 months was slightly higher in infants of depressed women compared to non-depressed women, this difference did not reach statistical significance. In line with our findings, a study involving 4745 children

Table 2: Anthropometric measurement according to postpartum depression at 6 and 12 months								
Measurement	Time							
	Depression status at 6 months		P	Depression status	P	Depression		
	No Depressed	Depressed		No Depressed	Depressed		status, P	
W-F-A, z-score	6.59 (0.30)	6.37 (0.37)	0.39	8.85 (0.36)	9.04 (0.44)	0.55	0.08	
Wfh, z-score	0.15 (0.31)	0.09 (0.38)	0.83	0.43 (0.33)	0.56 (0.40)	0.63	0.46	
Hfa, z-score	7.83 (0.40)	7.66 (0.49)	0.62	11.26 (0.49)	11.21 (0.60)	0.90	0.72	
Mfa, z-score	-0.07 (0.24)	-0.09 (0.29)	0.93	0.38 (0.24)	0.48 (0.30)	0.640	0.48	

Results were reported from repeated measures ANOVA. Data are shown as mean (SE). Wfh: weight-for height, W-F-A: weight-for-age, Hfa: height-for-age, and Mfa: MUAC-for-age

Table 3: Neurodevelopmental Skill measurement score according to postpartum depression at 6 and 12 months								
Measurement	Time							
	Depression status at 6 months of age		P	Depression status a	P	Depression		
	Not depressed	Depressed		Not depressed	Depressed		status, P	
Communication	53.02 (2.41)	53.55 (2.64)	0.64	54.99 (2.64)	53.47 (2.89)	0.22	0.02*	
Gross motor	58.91 (2.08)	58.10 (2.27)	0.41	58.93 (2.27)	57.40 (2.48)	0.14	0.27	
Fine motor	55.95 (1.95)	55.22 (2.13)	0.42	54.87 (1.91)	54.60 (2.09)	0.73	0.51	
Problem Solving	59.31 (1.67)	59.35 (1.83)	0.96	58.24 (1.86)	57.67 (2.03)	0.51	0.33	
Personal-Social	56.96 (2.84)	57.30 (3.11)	0.79	53.67 (3.34)	53.12 (3.65)	0.72	0.22	

<sup>\*</sup>P<0.05 is considered as statistically significant. Results were reported from repeated measures ANOVA. Data are shown as mean (SE)

CI 21 X7 - 2 - 1-2	Table 4: Association of p	ostpar tu			cai out velo	pment a			-
Skill Variables	Parameter	D.4.		odel a		D.4.		odel b	n
		Beta	95% CI		P	Beta	95% CI		P
Communication	Communication (6 months)	0.76	Lower	Upper 0.87	<0.001*	0.77	Lower 0.67	Upper 0.88	<0.001*
Communication (12 months)	` ,	0.76	0.66			0.77			
(12 months)	Gross movements (6 months)	0.06	-0.08	0.20	0.38	0.09	-0.05	0.23	0.19
	Fine movements (6 months)	-0.01	-0.16	0.14	0.90	0.02	-0.13	0.17	0.80
	Problem solving (6 months)	0.12	-0.05	0.28	0.16	0.13	-0.04	0.29	0.13
	Personal–Social (6 months)	0.04	-0.05	0.13	0.37	0.04	-0.05	0.13	0.43
	Depression	1.89	0.44	3.34	0.01*	1.81	0.32	3.30	0.02*
Gross	Communication (6 months)	-0.02	-0.11	0.06	0.60	-0.03	-0.12	0.06	0.53
movements (12 months)	Gross movements (6 months)	0.80	0.68	0.91	<0.001*	0.79	0.67	0.91	<0.001*
monuis)	Fine movements (6 months)	0.08	-0.05	0.20	0.23	0.07	-0.06	0.20	0.27
	Problem solving (6 months)	0.03	-0.10	0.17	0.61	0.04	-0.10	0.17	0.62
	Personal-Social (6 months)	0.01	-0.06	0.09	0.69	0.02	-0.06	0.10	0.60
	Depression	0.77	-0.42	1.95	0.20	0.47	-0.78	1.72	0.46
Fine	Communication (6 months)	0.03	-0.04	0.11	0.39	0.01	-0.06	0.09	0.74
movements (12	Gross movements (6 months)	0.06	-0.04	0.16	0.22	0.04	-0.07	0.14	0.49
months)	Fine movements (6 months)	0.65	0.54	0.76	<0.001*	0.63	0.52	0.74	<0.001*
	Problem solving (6 months)	0.01	-0.11	0.13	0.91	0.00	-0.12	0.12	0.96
	Personal-Social (6 months)	0.07	0.00	0.13	0.04*	0.07	0.00	0.13	0.04*
	Depression	0.13	-0.94	1.19	0.81	0.27	-0.79	1.34	0.62
Problem	Communication (6 months)	0.04	-0.03	0.11	0.27	0.00	-0.07	0.08	0.95
solving (12	Gross movements (6 months)	0.03	-0.07	0.12	0.58	0.01	-0.08	0.11	0.80
months)	Fine movements (6 months)	0.18	0.07	0.28	<0.001*	0.15	0.04	0.26	0.01*
	Problem solving (6 months)	0.66	0.55	0.77	<0.001*	0.65	0.54	0.77	< 0.001*
	Personal–Social (6 months)	0.05	-0.01	0.11	0.10	0.06	0.00	0.12	0.07
	Depression	0.40	-0.61	1.41	0.44	0.74	-0.29	1.76	0.16
Social	Communication (6 months)	0.03	-0.06	0.13	0.50	0.03	-0.07	0.13	0.55
Personality (12 months)	Gross movements (6 months)	0.03	-0.09	0.15	0.63	0.04	-0.09	0.16	0.56
	Fine movements (6 months)	-0.02	-0.16	0.12	0.77	0.01	-0.13	0.15	0.91
	Problem solving (6 months)	0.06	-0.09	0.21	0.42	0.06	-0.09	0.21	0.42
	Personal–Social (6 months)	1.01	0.93	1.09	<0.001*	1.03	0.94	1.11	<0.001*
	Depression	0.93	-0.39	2.24	0.17	1.09	-0.25	2.44	0.11

Model b adjusted for delivery type, maternal age at delivery, SES, number of previous children, maternal education, maternal employment, breastfeeding at 12 months, exclusive breastfeeding, supplementation, infant's gender, and birth weight. \*P<0.05 is considered statistically significant. Results were reported from the multivariate linear regression model

found no association between weight-for-length below the 10<sup>th</sup> percentile, while a significant association was observed with length, particularly in low/middle-income contexts.<sup>[35]</sup> However, contrary to earlier research, there was no confirmed increase in the likelihood of underweight or stunted growth among children of depressed mothers during the first year of life in developed countries.<sup>[12,36]</sup> This discrepancy may be attributed to similar rates of breastfeeding, exclusive breastfeeding, and gestational weight gain in depressed and non-depressed women in our study.

According to recent study findings, a decline and poor progression in communication skills from 6 to 12 months were observed, particularly at the 12 months. Other

studies have also reported disruptions in communication skills among infants of depressed mothers at various ages, ranging from 12 months<sup>[37,38]</sup> to 18 months.<sup>[39]</sup>

Furthermore, a longitudinal birth cohort study by Avan *et al.*<sup>[11]</sup> demonstrated a significant association between maternal postnatal depression and child behavior problems, including concentration difficulties and impaired social relationships, at the age of 2 years. Some studies have even suggested that children whose mothers suffer from PPD are at risk of emotional and psychological issues and delays in cognitive and language development.<sup>[9,15]</sup>

Additionally, the findings of a systematic review by Sohr-Preston and Scaramella suggest that late maternal depression (occurring between 5 and 12 weeks postpartum), but not early-onset maternal PPD (within 4 weeks after childbirth), was associated with reduced expressive language scores during infancy. Their study evaluated recipient and expressive communication skills at ages 10, 14, 18, 24, 32, and 40 months in infants with depressed mothers.<sup>[40]</sup>

Language development is a critical skill for effective communication, consisting of two facets: recipient and expressive language. Expressive language, which involves conveying a message, appears to be more indirectly influenced by maternal behavior than recipient skills, which encompasses receiving and comprehending messages from a caregiver. In line with our study, a descriptive investigation in Tehran involving 1036 mothers and their offspring has revealed delays in a child's communication skills when the mother experiences depression. [25]

A literature review supports the hypothesis that depressed mothers may manifest heightened irritability and diminished responsiveness to their infant's emotional and physical needs.<sup>[40]</sup>

Furthermore, interactions involving play, vocal communication, and explanations between depressed mothers and their infants tend to be minimized. [41,42] Additionally, parenting styles, including intrusive and passive behaviors, [42] insecure attachment, [14,43] reduced storytelling, and diminished verbal communication by depressed mothers, [44] expedite adverse effects on infant communication. Conversely, maternal depression can give rise to unhealthy behaviors that impact neurodevelopment.

In a study by Ibanez et al., [45] antenatal depression was correlated with children's development at 3 years of age, but only in the presence of antenatal anxiety. In other words, psychological disorders may significantly impact infants' neurodevelopmental delays more than maternal depression alone. No disparities were observed in other domains of the child's neurodevelopmental scores between depressed and non-depressed mothers. Nevertheless, study results indicated that the persistence of PPD would augment developmental delay in most<sup>[46]</sup> or some ASQ domains at 24 months. These discrepancies could be ascribed to methodological limitations and different assessment tools for infants' neurodevelopment, including the Bayley, ASQ, and Mullen Scales of Early Learning questionnaires. Furthermore, assessments of maternal depression were conducted at various time points.

Despite prior research findings, no definitive evidence has emerged concerning the specific domains of child neurodevelopmental delay. As mentioned previously, this inconsistency may be attributed to disparities in methodological approaches.

The present study possesses several strengths. Our data stem from a prospective birth cohort study that

accounts for various confounding factors. However, there are limitations to our research. It should be noted that depression levels were assessed only in the fourth week post-delivery, disregarding potential variations over time, and other influential factors affecting both infant birth size and PPD may have been overlooked.

#### **Conclusions**

The current study has elucidated a noteworthy correlation between Postpartum Depression (PPD), a prevalent health concern, and the neurodevelopmental scores of children. The evaluation of depression as a postpartum screening tool can prove to be a valuable asset for parents and caregivers, facilitating the identification of mothers who are susceptible and affording them early treatment and essential support services, thus alleviating potential detrimental impacts on their offspring.

#### **Abbreviations**

ASQ: Ages and Stages Questionnaires

BMI: Body Mass Index

EPDS: Edinburgh Postpartum Depression Scale

IBC: Isfahan Birth Cohort
MDI: Mental Development Index
OFC: Occipitofrontal Circumference

PPD: Postpartum Depression SES: Socioeconomic Status.

#### Declarations ethics approval and consent to participate

This study adhered to the principles of the Declaration of Helsinki and received approval from the Research Ethics Committees of the Medical University of Isfahan under the reference number IR.MUI.RESEARCH. REC.1398.331. All participants were duly informed that their participation was voluntary and that their responses would remain confidential. Informed written consent was obtained from all questionnaire respondents, and participants were assured they could withdraw from the study without incurring any penalties. No compensation was provided for participation in the relevant guidelines and regulations conducted in all methods and experiments.

#### Availability of data and materials

The corresponding authors are prepared to disclose the acquired data upon receiving reasonable requests.

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#### **Conflicts of interest**

There are no conflicts of interest.

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