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Association of Morningness-Eveningness with Psychiatric Symptoms Among Pregnant Women

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Abstract

In the present study, we examined associations between circadian preference and psychiatric symptoms among 1,796 pregnant from Lima, Peru. One quarter were classified as evening types. Compared to morning types, evening type pregnant women had increased odds of generalized anxiety (OR=1.44; 95% CI: 1.12–1.86) and posttraumatic stress disorder (OR=1.38; 95% CI: 1.07–1.78). Although there was a positive trend, evening chronotype was not significantly associated with elevated odds of depression (OR=1.23; 95% CI: 0.94–1.61). Future studies are warranted to help understand the underlying behavioral, biological, and genetic pathways of these associations. Assessing circadian preference may help clinicians identify pregnant women at risk for psychiatric symptoms.

Keywords

circadian rhythm; depression; anxiety; posttraumatic stress disorder; pregnancy

Introduction

Circadian dysregulation has been implicated in the pathogenesis of mood and other psychiatric disorders [1–3]. Notably, evening-types are more likely to report symptoms of psychiatric disorders [1], including depression [3, 4], anxiety [5, 6], bipolar disorder [7, 8], and posttraumatic stress disorder (PTSD) [9]. The literature concerning circadian preference and psychiatric disorders, however, is quite sparse for pregnant women, a population known to be particularly vulnerable to profound sleep disturbances [10–14]

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All authors declare no conflicts of interest.

and psychiatric disorders during gestation and the postpartum period [10, 15–17]. As psychological distress during pregnancy may increase the risk of adverse maternal and infant health outcomes, we sought to examine the association of circadian preference with symptoms of depression, anxiety, and PTSD in a cohort of well-characterized pregnant women enrolled in a prospective cohort study [18, 19].

Methods

Study population

This cross-sectional study is part of the Pregnancy Outcomes, Maternal and Infant Cohort Study (PrOMIS) conducted at the Instituto Nacional Materno Perinatal (INMP) in Lima, Peru between February 2012 and July 2014. Details of the study have been described previously [18, 20]. Briefly, PrOMIS was a longitudinal cohort study that examined social and behavioral maternal risk factors of pregnancy outcomes. Participants were pregnant women aged 18–47 who had their first prenatal care visit at the INMP. Women were eligible if they spoke and read Spanish, initiated prenatal care before 16 weeks' gestation, and planned to deliver at INMP. Women were excluded from the study if they were younger than 18 years of age or had completed more than 18 weeks' gestation. Though multiple interviews were conducted during the cohort, only information from the first interview was used for this study, which occurred during the first prenatal care visit (mean gestational age of 11 weeks). Participants were interviewed about sociodemographic and lifestyle characteristics, medical and reproductive histories, abuse history, and chronotype characteristics. All participants provided written informed consent and institutional review boards of the INMP, Lima, Peru, and the Harvard T.H. Chan School of Public Health, Office of Human Research Administration, Boston, MA, approved all procedures used in the study.

Munich Chronotype Questionnaire (MCTQ)

The Spanish-language Munich Chronotype Questionnaire was used to assess the sleep corrected mid-sleep on free days (MSFsc) [21]. The MSFsc assesses a participants' chronotype based on local time of mid-sleep on free days corrected for sleep debt accumulated during the workweek, excluding those who use alarm clocks or don't wake voluntarily free of disturbances. The MSFsc was calculated using the formula

 $MSF_{SC} = \frac{Midsleep \text{ on free days} - \binom{Sleep \text{ duration}}{on \text{ free days}} - \frac{average \text{ weekly}}{sleep \text{ duration}}}{2}$

The MSFsc calculation is consistent with what was done in previous studies [22, 23]. The MSFsc scores range between 0 and 12 with lower values corresponding to morningness and higher values corresponding to eveningness. Given the lack of established cutoffs, scores were binned into three tertiles. A total of 44.3% of women were in tertile 1 (morning), 31.3% tertile 2 (intermediate), and 24.6% tertile 3 (evening type).

Anxiety Symptoms (GAD-7)

Symptoms of generalized anxiety were assessed using a Spanish-language version of the Generalized Anxiety Disorder-7 Questionnaire (GAD-7) [24, 25]. The GAD-7 is a 7-item questionnaire that examines the following symptoms of generalized anxiety in the two-week period prior to screening: 1) nervousness; 2) inability to stop worrying; 3) excessive worry; 4) restlessness; 5) difficulty in relaxing; 6) easily irritated; and 7) fear of something awful happening. Participants were asked how often they were bothered by each of these symptoms with response categories of: "not at all," "several days," "more than half the days," and "nearly every day," scored from 0–3 for a total score of 0 to 21. A validation study performed on our cohort of Spanish-speaking pregnant women found the GAD-7 to have good reliability and validity. Based on this study, generalized anxiety was defined as a GAD total score 7 (sensitivity=73.3%; specificity=67.3%) [26]. The Cronbach's alpha, a measure of internal consistency, in the present study is 0.84.

PTSD Symptoms (PCL-C)

A Spanish-language version of the PTSD Checklist–Civilian (PCL-C) questionnaire was used to assess the symptomology and severity of PTSD during the month preceding the interview [27, 28]. The PCL-C consists of 17 items that assess three symptom clusters: re-experiencing the trauma, avoidance and numbing, and hyperarousal. Responses were scored on a 5-point Likert scale from "not at all bothersome" to "extremely bothersome." Total PCL-C score ranges from 17 to 85, with higher scores indicating a higher severity of PTSD symptoms. A PCL-C score 26 was used to indicate presence of PTSD symptoms in our population [29]. The PCL-C has been validated in our cohort of Spanish-speaking pregnant women [29]. The Cronbach's alpha in the present study is 0.88.

Depressive Symptoms (PHQ-9)

Symptoms of antepartum depression were assessed using the Spanish-language Patient Health Questionnaire-9 (PHQ-9) [30]. The PHQ-9 has been validated among Spanish-speaking pregnant populations [31, 32]. The PHQ-9 asks about depressive symptoms in the two weeks prior and assigns scores of 0–3 to the following responses: 'never,' 'several days,' 'more than half the days,' and 'nearly every day.' As a severity measure, the total score of PHQ-9 scores range from 0 to 27. Depression was defined as a score 10 [33]. The PHQ-9 has been validated in our cohort of Spanish-speaking pregnant women [31, 32] and has been found to have good reliability [34]. The Cronbach's alpha in the present study is 0.76.

Additional Covariates

Sociodemographic and pregnancy variables included were categorized as follows: age (18–19, 20–29, 30–34, and 35 years); maternal race (Mestizo vs. others); years of education (6, 7–12, and >12 years); marital status (married/living with partner vs. others); employment status (yes vs. no); difficulty accessing basic foods (hard vs. not very hard); parity (nulliparous vs. multiparous); planned current pregnancy (yes vs. no); smoke during pregnancy (yes vs. no); alcohol consumption before pregnancy (yes vs. no); gestational age at interview (in weeks); early pregnancy body mass index (BMI): underweight (<18.5

kg/m²), normal (18.5–24.9 kg/m²), overweight (25–29.9 kg/m²), obese (30 kg/m^2); and self-reported health in the last year (good vs. poor).

Statistical analysis

Multivariable logistic regression procedures were performed with chronotype as the main predictor variable and psychiatric symptoms (antepartum depression, generalized anxiety, PTSD) as the outcome variable. Potential *a priori* confounders were included in the models. Morning chronotype was used as the reference group. Statistical analyses were performed using IBM SPSS Statistics 21.0 (SPSS Inc, Chicago, Illinois).

Results

1796 participants were included in the analysis. Mean gestational age was 11 weeks. The majority of participants were of Mestizo ethnicity [mixed European and Indigenous ancestry] (80.0%), unemployed (52.1%), and had >12 years of education (55.0%). Difficulty of getting basics including foods was reported by 43.9% of participants. A total of 31.2% of women experienced symptoms of generalized anxiety, 30.6% experienced symptoms of PTSD, and 24.9% experienced symptoms of antepartum depression (See Table 1).

Among participants, the overall mean MSFsc was 2.67 ± 1.38 (range=0–23). The mean MSFsc for the morning types was 1.50 (SD=0.69), for intermediate types 3.0 (SD=0), and for evening types 4.36 (SD=1.17). Compared to morning types, evening types had increased odds of experiencing symptoms of generalized anxiety (OR=1.44; 95%CI:1.12–1.86) and PTSD (OR=1.38; 95%CI:1.07–1.78). However, evening types did not have significantly increased odds of experiencing symptoms of depression (OR=1.23; 95%CI:0.94–1.61) (See Table 2).

Discussion

Evening type pregnant women were more likely to experience symptoms of generalized anxiety and PTSD compared to morning types, while intermediate types did not have elevated odds of psychiatric disturbances. Morningness-eveningness was not associated with symptoms of depression.

While no previous studies have examined the association between morningness-eveningness and anxiety during pregnancy, this association has been studied among non-pregnant populations. Our findings are consistent with most [6, 35, 36] though not all [5, 23] prior studies that reported a positive association between eveningness and anxiety. We are aware of one previous study that examined the association between chronotype and PTSD. Hasler et al in their study among combat veterans [9] found a positive correlation between evening chronotype and PTSD (r=0.41). This is not surprising as one of the hallmarks of PTSD is a state of hyperarousal which is characterized by difficulty initiating and maintaining sleep [9].

While we found one study demonstrating higher odds of depression among morning types [35], the majority of previous studies have largely shown that evening types are more likely

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to report symptoms of depression [3, 5, 37–41]. Although a positive trend was present, we did not find a significant association between chronotype and depression in our population of pregnant women. There are a few possible explanations for our findings. Other studies show participants with depression are more likely to be intermediate than morning or evening types [42–44]. Furthermore, the prevalence of depression has been found to increase later in pregnancy [45] and the post-partum period [46], and this study assessed women at an average gestational age of 11 weeks. Another possible explanation is that sleep disturbances are common during pregnancy, which may alter the relationship between chronotype on mood [12, 47, 48].

Previous studies have demonstrated a connection between morningness-eveningness and psychological health. Mood disorders, including depression and bipolar disorder, are associated with circadian rhythm disruptions affecting sleep cycles, body temperature regulation, blood pressure, and secretion of cortisol and melatonin [49, 50]. Furthermore, reproductive hormones may alter rhythmicity and contribute to mood disturbances during pregnancy [51]. There is growing evidence that chronotypes are associated with the immune system and maintenance of biological systems during pregnancy [52]. Understanding chronotype disturbances during pregnancy is important given the association between psychiatric disorders and poor infant neurocognitive and physical development outcomes.

Our study expands the literature by assessing the association between morningnesseveningness and psychiatric symptoms among pregnant Peruvian women. The strengths of our study include a large sample size and use of previously validated questionnaires. However, our study has some limitations. First, the study was conducted at a single hospital in Lima, Peru, which primarily serves a low-income population, and our results may not be generalizable to other pregnant populations. Second, generalized anxiety, PTSD, and depression were assessed based on screening instruments. Although the use of brief screening instruments is the most feasible method of data collection for large-scale epidemiologic studies, future studies using diagnostic evaluations for mood disorders are warranted.

Overall, our findings suggest that circadian preference has important implications for psychiatric disorders during pregnancy. When compared to morning types, evening type pregnant women have significantly increased odds of experiencing generalized anxiety and PTSD symptoms. As the prevalence and severity of psychiatric disorders vary during pregnancy, longitudinal studies are warranted to help understand how circadian preference throughout pregnancy may impact psychiatric symptoms. Assessing morningness-eveningness may help clinicians identify pregnant women at increased risk of experiencing symptoms of anxiety and PTSD during pregnancy and may also contribute to our understanding of the impact of circadian rhythm on the overall health of pregnant women.

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Data availability statement

Data is available upon request.

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Characteristics	All participants (N = 1,796)		Morning type: (N= 796)		Intermediate type: (N= 558)		Evening type: (N= 442)		p-value
	n	%	n	%	n	%	n	%	
Age (years) ^{a}	27	$.77 \pm 6.25$	28	.76 ± 6.43	2'	7.86 ± 6.12	25	.86 ± 5.65	<0.001
Age (years)									
18–20	118	6.6	46	5.8	32	5.7	40	9.0	<0.001
20–29	1010	56.2	406	51.0	319	57.2	285	64.5	
30–34	377	21.0	177	22.2	120	21.5	80	18.1	
35	291	16.2	167	21.0	87	15.6	37	8.4	
Early pregnancy BMI (kg/ m ²) ^{<i>a</i>}	25.68 ± 4.18		25.90 ± 4.32		25.50 ± 4.11		25.49 ± 3.99		0.155
Early pregnancy BMI (kg/m ²)									
<18.5	36	2.0	15	1.9	12	2.2	9	2.0	0.394
18.5–24.9	818	45.5	347	43.6	256	45.9	215	48.6	
25–29.9	676	37.6	298	37.4	217	38.9	161	36.4	
30	248	13.8	125	15.7	68	12.2	55	12.4	
Gestational age at interview ^a	11.17 ± 3.93		11.03 ± 3.91		11.30 ± 3.96		11.27 ± 3.95		0.341
Education (years)									
6	22	1.2	12	1.5	6	1.1	4	0.9	0.007
7–12	779	43.4	380	47.7	231	41.4	168	38.0	
>12	987	55.0	402	50.5	316	56.6	269	60.9	
Mestizo ethnicity	1437	80.0	645	81.0	440	78.9	352	79.6	0.781
Married/living with a partner	1479	82.4	657	82.5	474	85.3	348	78.7	0.027
Employed	861	47.9	421	52.9	249	44.6	191	43.2	0.001
Difficulty accessing basic food	786	43.9	351	44.3	261	47.0	174	39.4	0.051
Difficulty accessing medical care	740	41.5	347	43.9	224	40.8	169	38.3	0.152
Nulliparous	836	46.6	342	43.0	247	44.3	247	55.9	<0.001
Planned pregnancy	690	38.4	303	38.1	218	39.1	169	38.2	0.800
Tobacco smoking before pregnancy	265	14.8	89	11.2	75	13.4	101	22.9	<0.001
Tobacco smoking during pregnancy	33	1.8	13	1.6	5	0.9	15	3.4	0.026
Alcohol consumption before pregnancy	574	32.0	211	26.5	181	32.4	182	41.2	<0.001
Alcohol consumption during pregnancy	93	5.2	29	3.6	32	5.7	32	7.2	0.053
Antepartum generalized	5	$.15 \pm 3.84$	4	$.92 \pm 3.81$:	5.00 ± 3.59	5	$.75 \pm 4.14$	0.001

Characteristics of population according to circadian preference in PrOMIS 2 (N = 1,796)

anxiety^a

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Characteristics	All participants (N = 1,796)		Morning type: (N= 796)		Intermediate type: (N= 558)		Evening type: (N= 442)		p-value
	n	%	n	%	n	%	n	%	•
Antepartum generalized anxiety (GAD 7)	560	31.2	230	28.9	174	31.2	156	35.3	0.066
Antepartum PTSD ^a	23.63 ± 7.69		23.53 ± 7.52		23.15 ± 7.06		24.41 ± 8.64		0.035
Antepartum PTSD (PCL 26)	550	30.6	228	28.6	166	29.7	156	35.3	0.045
Antepartum depression ^a	6.92 ± 4.36		6.69 ± 4.26		6.92 ± 4.25		7.34 ± 4.65		0.037
Antepartum depression (PHQ-9 10)	448	24.9	191	24.0	138	24.7	119	26.9	0.517

Due to missing data, percentages may not add up to 100%. For continuous variables, P-value was calculated using ANOVA; for categorical values, P-value was calculated using Chi-square test. Abbreviations: BMI; Body Mass Index.

^{*a*}Mean \pm standard deviation

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Table 2.

Association of circadian preference with psychiatric symptoms during pregnancy

	No psychiatric symptoms		Psychiatric symptoms					
	Ν	%	Ν	%	Unadjusted OR (95% CI)	Adjusted OR (95% CI) ^a		
Antepartum generaliz	zed anxiety							
Circadian preference								
Morning	566	45.8	230	41.1	Reference	Reference		
Intermediate	384	31.1	174	31.1	1.12 (0.88–1.41)	1.14 (0.90–1.45)		
Evening	286	23.1	156	27.9	1.34 (1.05–1.72)	1.44 (1.12–1.86)		
p-value for trend					0.022	0.005		
Antepartum PTSD								
Circadian preference								
Morning	568	45.6	228	41.5	Reference	Reference		
Intermediate	392	31.5	166	30.2	1.06 (1.83–1.34)	1.04 (0.82–1.32)		
Evening	286	23.0	156	28.4	1.36 (1.06–1.74)	1.38 (1.07–1.78)		
p-value for trend					0.021	0.014		
Antepartum depressi	on							
Circadian preference								
Morning	605	44.9	191	42.6	Reference	Reference		
Intermediate	420	31.2	138	30.8	1.04 (0.81–1.34)	1.04 (0.81–1.35)		
Evening	323	24.0	119	26.6	1.17 (0.90–1.52)	1.23 (0.94–1.61)		
p-value for trend					0.270	0.145		

Abbreviations: OR, odds ratio; CI, confidence interval.

^aAdjusted for age (continuous), marital status (married vs. others), parity (nulliparous vs. others), difficulty paying for the very basics (hard vs. not very hard)