Reliability and validity of the PHQ-8 in first-time mothers who used assisted reproductive technology

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STUDY QUESTION: Is the Patient Health Questionnaire-8 (PHQ-8) a valid and reliable measure of depression in first-time mothers who conceived via ART?

SUMMARY ANSWER: The results from this study provide initial support for the reliability and validity of the PHQ-8 as a measure of depression in mothers who have conceived using ART.

WHAT IS KNOWN ALREADY: Women who achieved a clinical pregnancy using ART experience many stressors and may be at an increased risk of depression. The PHQ-8 is a brief measure designed to detect the presence and severity of depressive symptoms. It has been validated in many populations; however, it has not been validated for use in this population.

STUDY DESIGN, SIZE, DURATION: This is a cross-sectional study of 171 first-time mothers in the USA, recruited through Amazon’s Mechanical Turk (MTurk).

PARTICIPANTS/MATERIALS, SETTING, METHODS: The reliability of the PHQ-8 was measured through a Cronbach’s alpha, the convergent validity was measured through the correlation between the PHQ-8 and the Generalized Anxiety Disorder-7 (GAD-7) measure of anxiety symptoms, and the structural validity was measured through a Confirmatory Factor Analysis.

MAIN RESULTS AND THE ROLE OF CHANCE: The Cronbach’s alpha for the total PHQ-8 was acceptable (α = 0.922). The correlation between the PHQ-8 and the GAD-7 was large (r = 0.88) indicating good convergent validity. Ultimately, a bifactor model provided the best model fit (χ²(13) = 23.8, P = 0.033; Comparative Fit Index = 0.987; Root Mean Square Error of Approximation = 0.07, Tucker–Lewis Index = 0.972).

LIMITATIONS, REASONS FOR CAUTION: The results are limited by: the predominantly white and well-educated sample, a lack of causation between the use of artificial reproductive technology and depressive symptoms, including mothers with children up to 5 years old, convergent validity being based on associations with a related construct instead of the same construct, lack of test-retest reliability, divergent validity and criterion-related validity, data collected through MTurk, and the fact that the measures used were all self-report and therefore may be prone to bias.

WIDER IMPLICATIONS OF THE FINDINGS: Consistent with previous literature, a bifactor model for the PHQ-8 was supported. As such, when assessing depression in first-time mothers who conceived via ART, using both the PHQ-8 total score and subdomain scores may yield the most valuable information. The results from this study provide preliminary support for the reliability and validity of the PHQ-8 as a measure of depression in first-time mothers who conceived using ART.

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**WHAT DOES THIS MEAN FOR PATIENTS?**

Due to the many stressors that face first-time mothers who conceive using infertility treatments, they may be at risk for depression. While they may be at high risk, little is known about whether our existing measures of depression behave the same way in this population.

This study examines a common measure of depression (the Patient Health Questionnaire-8 or PHQ-8) to see if it accurately and reliably captures depressive symptoms in first-time mothers who conceived with the use of infertility treatments.

After examining the results of the statistical processes, we are able to provide support for the use of the PHQ-8 in this population. Furthermore, we provide support for breaking the PHQ-8 into two subscales to provide valuable information for this population.

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**Introduction**

The International Committee for Monitoring Assisted Reproductive Technologies (ICMART) defines infertility as a disease that is characterized by the failure to establish a clinical pregnancy after 12 months of regular, unprotected sexual intercourse or an impairment of a person's capacity to reproduce either as an individual or with his/her partner (ICMART, 2017). Rates of infertility are rising with one in six couples who want to conceive being diagnosed with infertility (Ravitsky and Kimmins, 2019). ART, which is defined as all interventions that involve the *in vitro* handling of both human oocytes and sperm or of embryos for the purpose of reproduction (ICMART, 2017), increases the likelihood of achieving a clinical pregnancy in couples experiencing infertility. Nonetheless, these treatments are financially costly (Collins, 2001; Katz et al., 2011) and physically and psychologically burdensome (Aimagambetova et al., 2020) especially for females as ART procedures (i.e. daily shots, hormone treatments, egg retrieval, embryo transfer) are largely performed on the woman. Women who have achieved a clinical pregnancy using ART may be at an increased risk for experiencing depression (Ross et al., 2011; Gdańska et al., 2017). This heightened risk for depression and avoidance of negative feelings may continue during the transition to parenthood especially for first-time mothers who may be more likely to idealize parenthood, experience greater concerns about their child’s health, and feel less entitled to seek social support when they feel doubts or uncertainty about parenting (Ulrich et al., 2005; Gressier, 2015). Furthermore, continued infertility and challenges to conceive subsequent children naturally may have a negative impact on the psychological well-being of mothers after conceiving via ART (Hjelmstedt et al., 2004). As such, it is important to have psychometrically sound measures than can assess depressive symptoms in mothers who have conceived using ART, particularly during the transition to parenthood.

The eight-item Patient Health Questionnaire (PHQ-8) is a brief measure designed to detect the presence and severity of depressive symptoms in adults (Kroenke et al., 2002). This measure has demonstrated acceptable internal consistency reliability, test–retest reliability, construct validity, factorial invariance and concurrent validity in Mexican and Central American descent university students residing in the USA, adults from Sweden with Systematic Sclerosis, and Latino/a university students living in the USA (Alpizar et al., 2018a,b; Mattsson et al., 2020). To the best of our knowledge, the psychometric properties of the PHQ-8 have not been previously evaluated in a sample of mothers who conceived using ART. Given increasing rates of infertility (Ravitsky and Kimmins, 2019) and the potential of a greater propensity for depression among first-time mothers who conceived via ART during the transition to parenthood (Ross et al., 2011; Gdańska et al., 2017), the current study sought to evaluate the reliability and validity of the PHQ-8 in first-time mothers of children 5 years old or younger who conceived using ART.

**Materials and methods**

**Participants**

The data used in this study were collected as a part of a larger study focused on assessing differences in maternal ratings of child vulnerability between first-time mothers who conceived using ART versus spontaneous conception (Egan et al., 2021). For the current study focused on assessing the psychometric properties of the PHQ-8 in first-time mothers who conceived using ART, only mothers who used ART were included. The sample consisted of 171 first-time mothers. Participants met inclusion criteria if they lived in the USA, were at least 18 years old, a first-time mother of a singleton child 5 years old or younger, endorsed experiencing infertility (which was defined as a failure to attain a clinical pregnancy after 12 months or more of trying to conceive), and reported utilizing a form of ART (i.e. IVF, ICSI, donor egg IVF, gestational carrier IVF, intrauterine embryo implantation, frozen embryo transfer, gamete intrafallopian transfer or zygote intrafallopian transfer) that resulted in the live birth of their child. Mothers of children up to 5 years old were included due to research suggesting that the effects of infertility and ART are far reaching and long lasting (Schmidt, 2010).

**Procedures**

Participants for this study were recruited during Spring 2018 using Amazon’s Mechanical Turk (MTurk). To ensure data quality, the study was advertised as one about parenting and conception methods and a screening survey was administered (Chandler and Shapiro, 2016; Thomas and Clifford, 2017). Once study eligibility was determined from the screening survey, an online consent form was presented to the participant. Participants who provided their online consent to participate in the study were offered the full set of questionnaires including the PHQ-8 and the Generalized Anxiety Disorder-7 (GAD-7).

After completing the survey, each participant was assigned a unique code to verify their participation through Qualtrics and receive compensation through MTurk. Participants received $1.81 for participation in the study and were not allowed to participate more than once.
Ethical approval
The study procedures outlined above were approved by the authors’ Institutional Review Board (IRB) ID#1395596-1.

Measures
Depression
Self-reported depression was measured by the PHQ-8 (Kroenke et al., 2009). The 9-item Patient Health Questionnaire (PHQ-9), from which the PHQ-8 is derived, is a widely used assessment for presence and severity of depressive symptoms (Kroenke et al., 2002). One item included on the PHQ-9 assesses suicide ideation and due to the inability of researchers to adequately respond to de-identified participants reporting suicide ideation, the PHQ-8 was developed with this item excluded (Kroenke et al., 2009). Exclusion of this item does not influence the sensitivity of the measure in detecting major depression (Kroenke et al., 2009). The PHQ-8 has been validated as a diagnostic tool and measure of depressive symptoms in clinical settings and large surveys in populations (Kroenke and Spitzer, 2002; Kroenke et al., 2009).

Participants were asked to reflect on their past 2 weeks and respond to 8 items on a 4-point Likert-type scale ranging from ‘not at all’ (0) to ‘nearly every day’ (3). Example items include: ‘little interest or pleasure in doing things’, ‘feeling tired or having little energy’ and ‘feeling down, depressed or hopeless’. Item responses were subsequently totaled and measured on a scale ranging from 0 to 24. Scores were interpreted as follows: 0–4 (minimal/no depression), 5–9 (minimal depression), 10–14 (moderate depression), 15–19 (moderate-severe depression), 20–24 (severe depression) (Kroenke and Spitzer, 2002; Kroenke et al., 2009, 2010).

Anxiety
Self-reported anxiety was measured by the GAD-7 (Spitzer et al., 2006). This 7-item assessment has been validated to measure a unidimensional factor of general anxiety disorder in the general population (Lowe et al., 2008; Naeinian et al., 2011). Participants were asked to reflect upon the last 2 weeks and answer, on a Likert-type scale ranging from ‘not at all’ (0) to ‘nearly every day’ (3), how often they had been bothered by the following problems: (i) feeling nervous, anxious, or on edge, (ii) not being able to stop or control worrying, (iii) worrying too much about different things, (iv) trouble relaxing, (v) being so restless that it is hard to sit still, (vi) becoming easily annoyed or irritable and (vii) feeling afraid, as if something awful might happen. Item responses were subsequently totaled and measured on a scale ranging from 0 to 21. Scores were interpreted as follows: 0–4 (minimal anxiety), 5–9 (mild anxiety), 10–14 (moderate anxiety), 15–21 (severe anxiety) (Spitzer et al., 2006).

The GAD-7 has been used to support convergent validity in previous validation studies of measures of depression (Lowe et al., 2008). Specifically, the GAD-7 and the PHQ-9 have been shown to be strongly correlated (Quon et al., 2015; Sawaya et al., 2016; Peters et al., 2021; Sequeira et al., 2021). Anxiety has been previously shown to be significantly correlated with depression in women who have conceived using ART (Huang et al., 2019). Therefore, in order to assess convergent validity through the measurement of an associated construct, the GAD-7 was included in this study.

Demographic questionnaire
Participants were asked to respond to a questionnaire assessing the following information: maternal age, age of first child, income, education, employment, maternal age at child’s birth, incidence of miscarriage, whether or not their child was born prematurely, marital status, number of ART treatments, cause of infertility, whether or not their insurance covered their ART treatments and what type of ART they used to conceive their child.

Statistical analysis
IBM Statistical Package for the Social Sciences (SPSS) Version 26 was used for this project. Statistical significance was determined by a P-value <0.05. An examination of skewness was conducted for the PHQ-8 and the GAD-7. Due to the nature of MTurk, there were no missing data and no data were excluded.

Floor and ceiling effects
An examination of floor and ceiling effects was conducted for the PHQ-8. Floor and ceiling effects were determined by examining whether or not greater than 15% of participants received either the lowest (floor) or highest (ceiling) possible score (McHorney and Tarlov, 1995; Terwee et al., 2007). Results with floor or ceiling effects indicate potentially poor content validity (Terwee et al., 2007).

Internal consistency reliability
Internal consistency reliability was assessed by examining Cronbach’s alphas for the PHQ-8. An acceptable range for Cronbach’s alpha is a value of 0.70 or higher (Nunnally, 1978). Inter-item correlations and the modified Cronbach’s alpha associated with the deletion of each item were also examined. Correlations were considered small $r \leq 0.1$, medium $r \geq 0.3$ and large $r \geq 0.5$ (Cohen, 1988).

Convergent validity
Convergent validity may be defined as the magnitude of the zero-order correlation between two closely related measures (Carlson and Herdman, 2012). To assess convergent validity, Pearson correlations were examined between the PHQ-8 and the GAD-7 (Spitzer et al., 2006). Correlations were considered small $r \leq 0.1$, medium $r \geq 0.3$ and large $r \geq 0.5$ (Cohen, 1988).

Structural validity
Structural validity of the PHQ-8 was assessed through a Confirmatory Factor Analysis (CFA) using R version 3.6.1. There is evidence from a Monte Carlo Simulation to suggest that for a multiple-factor model with 6–8 indicators, the minimum sample size needed is 100 participants (Wolf et al., 2013). Therefore, the sample included in this study of 171 was sufficient. The R code for the CFA can be found in the Supplementary Information. The development literature for the PHQ-8 suggests that the scale is measuring one factor (Kroenke et al., 2009). However, previous literature has suggested that in some populations, a two-factor model may present a more accurate model fit (Mattsson et al., 2020). Furthermore, there is evidence from work done with both the PHQ-8 and PHQ-9 that suggests a bifactor model, which estimated model fit based on a general factor of depression as well as two latent variables, is the superior model (Doi et al., 2018; Dong et al., 2019; Fischer et al., 2021). Therefore, this study examined a single factor, a two-factor, and a bifactor model. For both the two-factor model
and bifactor model, the two latent variables specified were cognitive/affective aspects of depression (items 1, 2, 6 and 7) and somatic aspects of depression (items 3, 4, 5 and 8) (Mattsson et al., 2020).

Within the CFA, the chi-square statistic (Hu and Bentler, 1999) was examined along with other measures of model fit including the Root Mean Squared Error of Approximation (RMSEA) Comparative Fit Index (CFI) and Tucker–Lewis index (TLI). In order to achieve excellent model fit, RMSEA values must be equal to or less than 0.06 and in order to achieve acceptable model fit, RMSEA values should be <0.08 (Browne and Cudeck, 1992; Hu and Bentler, 1999). In order to achieve excellent model fit, CFI and TLI values must be equal to or greater than 0.95 and in order to achieve acceptable fit, CFI and TLI values should range between 0.90 and 0.95 (Mulaik et al., 1989; Bentler, 1990; Hu and Bentler, 1995).

Results

Demographic data

Socio-demographic characteristics for this sample are provided in Table I. Most participants were around 30 years old (SD = 4.65, Range = 22–46) and were about 28 years old at the time of the birth of their first child (SD = 4.65, Range = 20–44). The majority of participants were married (86.5%), had at least a 4-year degree (72.5%), were employed (86%), and had children over the age of 18 months (65.5%). A large portion of participants also reported an income at or above $35 000 (58.5%), were White (76%), experienced infertility caused by a female factor (56.7%), had experienced one or more miscarriages (49.7%), did not have their child prematurely (76.4%) and received at least some financial help with infertility treatments from insurance (80.1%). The most common form of ART used was IVF with 73.3% of participants reporting having used IVF at some point throughout their infertility treatments. About 49% of women in this sample had undergone one to three cycles of ART. In this sample, 36.4% of mothers reported moderate to severe depressive symptoms.

### Table I Demographic variables for the sample.

| Characteristic                        | N or mean | % or SD | Range   |
|---------------------------------------|-----------|---------|---------|
| Age                                   | 30.37     | 4.65    | 22–46   |
| Age of eldest child                   | 1.92      | 1.30    | 0–5     |
| Age at birth of first child           | 28.02     | 4.63    | 20–44   |
| Marital status                        |           |         |         |
| Married                               | 148       | 86.5%   | –       |
| Divorced                              | 2         | 1.2%    | –       |
| Single                                | 20        | 11.7%   | –       |
| Separated                             | 1         | 0.6%    | –       |
| Cause of infertility                  |           |         |         |
| Male-factor                           | 33        | 19.3%   | –       |
| Female-factor                         | 97        | 56.7%   | –       |
| Both                                  | 10        | 5.8%    | –       |
| No known cause                        | 31        | 18.1%   | –       |
| Previous instances of miscarriage     |           |         |         |
| 0                                     | 86        | 50.3%   | –       |
| 1                                     | 53        | 31.0%   | –       |
| 2                                     | 25        | 14.6%   | –       |
| 3                                     | 6         | 3.5%    | –       |
| 4                                     | 1         | 0.6%    | –       |
| Was your child born prematurely?       |           |         |         |
| Yes                                   | 40        | 23.4%   | –       |
| No                                    | 131       | 76.36%  | –       |
| Did insurance cover your ART treatment cycles? |         |         |         |
| No                                    | 34        | 19.9%   | –       |
| Yes, partially                        | 83        | 48.5%   | –       |
| Yes, fully                            | 54        | 31.6%   | –       |

(continued)
Floor and ceiling effects
Floor effects on the PHQ-8 occurred in 15.8% of participants; 0.6% of participants reported a ceiling effect on the PHQ-8 total score. For the cognitive/affective subscale of the PHQ-8, floor effects occurred in 31.6% of the population. Ceiling effects occurred in 0.6% of participants on the cognitive/affective subscale of the PHQ-8. For the somatic subscale of the PHQ-8, floor effects occurred in 18.1% of participants. Ceiling effects occurred in 0.6% of the sample on the somatic subscale of the PHQ-8.

Internal reliability
Cronbach’s alpha for the PHQ-8 total score in this sample was within the acceptable range (α = 0.922). The Cronbach’s alphas for the cognitive/affective and the somatic subscales were also within the acceptable range (α = 0.867, α = 0.850, respectively). Table II presents the inter-item correlations for each of the eight items in the scale along with the Cronbach’s alpha if that item were to be deleted. All of the items correlated highly with each other (r’s ranged from 0.47–0.71). The Cronbach’s alphas would decrease with the deletion of any item in the scale.

Convergent validity
In support of convergent validity, correlation between the PHQ-8 and the GAD-7 was in the large range (GAD-7 r = 0.88, P < 0.001). Similarly, the cognitive/affective and the somatic subscales of the PHQ-8 correlated strongly with measures of maternal anxiety (cognitive/affective: GAD-7 r = 0.873, P < 0.001; somatic: GAD-7 r = 0.811, P < 0.001).

Structural validity
The skewness of each item was assessed and found to fall within the normal distribution. Fit indices for the CFAs can be found in Table III. Item loadings can be found in Table IV. The CFA testing a one-factor model used maximum-likelihood estimators. The one-factor model demonstrated adequate to excellent fit on most indices, $\chi^2(20) = 52.83, P < 0.001$; CFI = 0.961; RMSEA = 0.098; TLI = 0.945. All items loaded significantly onto the latent factor (β > 0.82).

Discussion
The purpose of the present study was to evaluate the psychometric properties of the PHQ-8 in mothers who conceived using ART. Women who have conceived via ART are at an increased risk for experiencing emotional distress (Aimagambetova et al., 2020). Consistent with previous research (Drosdzol and Skrzypulec, 2009;...
scores on the GAD-7 which assess maternal anxiety symptoms. The PHQ-8 via ART, using both the PHQ-8 total score and subdomain scores when assessing depression in first-time mothers who conceived may yield the most valuable information. Furthermore, the PHQ-8 model demonstrated the best overall fit in our sample. As such, models for the PHQ-8 (Alpizar, 2018a, b) consistent with previous literature testing single factor and two-factor model was a better fit than a two-factor model. This result is consistent with previous literature testing single factor and two-factor models for the PHQ-8 (Alpizar et al., 2018a, b). The bifactor model demonstrated the best overall fit in our sample. As such, when assessing depression in first-time mothers who conceived via ART, using both the PHQ-8 total score and subdomain scores may yield the most valuable information. Furthermore, the PHQ-8 total score and subdomain scores were highly correlated with scores on the GAD-7 which assess maternal anxiety symptoms. Since depression and anxiety are often comorbid (Spitzer et al., 2006) this indicates strong convergent validity ($r = 0.88$, $P < 0.001$). Taken as a whole, data from this study provide preliminary support for utilization of the PHQ-8 as a measure of depression in first-time mothers who have conceived using ART.

This study had a number of limitations. Given that the sample was predominantly white and well-educated, the present findings may not generalize to more diverse mothers who conceived using ART. We were not able to establish in our study that maternal depressive symptoms were a result of the ART experience; it would have been beneficial to include in the study a measure of stressful life events in order to control for other situations that may have been affecting maternal adjustment. Furthermore, our sample was comprised of first-time mothers who had a child 5 years old or younger and it is possible that, given this wide age range of children, mothers in our sample experienced different types of stressors that impacted their psychological well-being. It should be noted that we computed post hoc a Pearson correlation between maternal PHQ-8 scores and the age of child; this correlation ($r = −0.14$) was small and not statistically significant ($P = 0.066$), indicating child age may have had a small impact on maternal ratings of their depressive symptoms. Additionally, our assessment of convergent validity was based on a measure of a theoretically related construct, not by measuring the same construct. We also were not able to assess test-retest reliability, divergent validity, and criterion-related validity of the PHQ-8. Another limitation of this study was collecting data via Mechanical Turk. While Mechanical Turk has been shown to yield quality data (Kees et al., 2017) and participants in our study answered screening questions, there was no way for us to objectively determine maternal utilization of ART to conceive. Finally, our study relied entirely on maternal self-reports which may be prone to bias. Future research should focus on assessing test–retest reliability, divergent validity, and criterion-related validity of the PHQ-8 in mothers who have conceived via ART and there would be merit.

| Indicator | One-factor model | Two-factor model | Bifactor model |
|-----------|------------------|------------------|---------------|
|           | Cog./Aff. Somatic | Cog./Aff. Somatic | g             |
| Item 1    | 1.000            | 1.000            | 1.000         |
| Item 2    | 1.030            | 1.030            | 1.322         |
| Item 3    | 0.911            | 1.000            | 1.000         |
| Item 4    | 0.819            | 0.899            | 1.018         |
| Item 5    | 0.981            | 1.077            | 0.367         |
| Item 6    | 1.024            | 1.024            | 0.574         |
| Item 7    | 0.964            | 0.964            | 0.359         |
| Item 8    | 0.996            | 1.094            | −1.243        |

Item 1: Little interest or pleasure in doing things; Item 2: Feeling down, depressed or hopeless; Item 3: Trouble falling or staying asleep, or sleeping too much; Item 4: Feeling tired or having little energy; Item 5: Poor appetite or overeating; Item 6: Feeling bad about yourself or that you are a failure or have let yourself or your family down; Item 7: Trouble concentrating on things such as reading the newspaper or watching television; Item 8: Moving or speaking so slowly that other people could have noticed—or the opposite—so fidgety or restless that you have been moving around a lot more than usual. PHQ-8, Patient Health Questionnaire-8; Cog./Aff., Cognitive/Affective; g, effect size Hedges-g.
in reproducing this study in an in-person setting rather than online.

In conclusion, the results from this study provide preliminary support for the reliability and validity of the PHQ-8 as a measure of depression in first-time mothers who conceived using ART.

Supplementary data
Supplementary data are available at Human Reproduction Open online.

Data availability
The data underlying this article cannot be shared publicly due to the outlined agreement with the IRB at the author’s institution. The data will be shared on reasonable request to the corresponding author.

Authors’ roles
C.P., K.E. and C.L.: (i) substantial contributions to conception and design, or acquisition of data, or analysis and interpretation of data, (ii) drafting the article or revising it critically for important intellectual content, (iii) final approval of the version to be published and (iv) agreement to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

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Conflict of interest
The authors declare that they have no conflicts of interest.

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