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Prenatal electrocardiogram testing and postpartum depression: A population-based cohort study

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Abstract

Background: Cardiovascular symptoms in pregnancy may be a clue to psychological distress. We examined whether electrocardiogram testing in pregnant women is associated with an increased risk of subsequent postpartum depression.

Methods: We conducted a population-based cohort study of pregnant women who delivered in Ontario, Canada comparing women who received a prenatal ECG to women who did not.

Results: In total, 3,238,218 women gave birth during the 25-year study period of whom 157,352 (5%) received an electrocardiogram during prenatal care. Receiving an electrocardiogram test was associated with a one-third relative increase in the odds of postpartum depression (odds ratio 1.34; 95% confidence interval 1.29–1.39, p < 0.001).

Conclusion: The association between prenatal electrocardiogram testing and postpartum depression suggests a possible link of organic disease with mental illness, and emphasizes that cardiovascular symptoms may be a clinical clue to the presence of an underlying mood disorder.

Keywords

Depression, postpartum, pregnancy complications, cardiovascular, cardiovascular testing

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Introduction

Postpartum depression affects 1 in 10 women in the year following childbirth.^{1,2} Classic features include decreased mood, disturbances of sleep, excessive fatigue, weight changes, and feelings of worthlessness that can be particularly problematic for new mothers who are adjusting to life with a new baby.^{3–5} Postpartum depression can interfere with breastfeeding, impair infant bonding, and potentially effect neurocognitive development for a child.^{6–8} In 2017, the total societal cost of untreated perinatal mood and anxiety disorders in the US was \$14.2 billion.⁹ In addition, suicide is a major cause of postpartum mortality with rates of 1 to 5 per 100,000 in high income countries.^{10–12}

Electrocardiograms (ECGs) are simple diagnostic tests that pose no real immediate physical risks to a pregnant woman. Guidelines suggest ECGs can be indicated for pregnant women presenting with chest pain, shortness of breath, or unexplained palpitations.¹³ However, such symptoms are often due to normal physiologic changes of pregnancy rather than serious pathology. The heart rate increases substantially during pregnancy and ectopic beats with a non-sustained arrhythmia are detected in over half of pregnant women.¹⁴ Furthermore, one-in-seven pregnant women have nonspecific ECG waveform changes that completely resolve following delivery.¹⁵ These ECG abnormalities rarely indicate cardiovascular disease.

Mood disorders frequently manifest with somatic symptoms, and women with depression have a high prevalence of sinus arrhythmias.¹⁶ In addition, simple medical tests may be distressing to some women, particularly if results are equivocal and more investigations are pursued.¹⁷ In the general adult population, ECG testing is associated with a fivefold risk of additional cardiovascular procedures.¹⁸ Moreover, pregnant women may be less accustomed to extensive medical testing and ECGs are not part of routine prenatal assessments. Whether cardiovascular testing in pregnancy can predict postpartum depression is unknown. We examined whether ECG testing of women during pregnancy was associated with an increased risk of subsequent postpartum depression.

Methods

Study setting

We performed a population-based cohort study of postpartum women in Ontario, Canada to examine the association between ECG testing in pregnancy and postpartum depression. Ontario is

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Donald A Redelmeier, Sunnybrook Health Sciences Centre, G-151, 2075 Bayview Ave, Toronto, Ontario, Canada M4N 3M5. Email: dar@ices.on.ca Canada's largest province with a population of 12,160,282 and 135,595 live births in 2006 (study midpoint).^{19,20} In Ontario, women have access to universal health insurance coverage with no out-of-pocket costs for medical services including prenatal care, ECG testing, and psychiatric services. In addition, health care records can be tracked through established database analyses.²¹ The study protocol was approved by the Research Ethics Board at Sunnybrook Health Sciences Centre, including a waiver for individual's consent.

Patient identification

We identified women 14–55 years old who had a live birth in Ontario, Canada between 1 April 1993 and 31 March 2018 (enrolment interval of 25 years). This interval was selected to cover all available data and provide a minimum follow-up of one-year after delivery. To identify eligible women we linked maternal–newborn pairs using the MOMBABY database, which identified more than 98% of all Ontario births.²² We excluded women with a missing health card as they were ineligible for healthcare coverage under the Ontario Health Insurance Plan (OHIP). Different pregnancies (by the same mother) were coded as separate occurrences so that each delivery was included for each woman (no statistical adjustments for possible clustering). Pregnancies resulting in a stillbirth were excluded.

Electrocardiogram testing

The primary analysis compared women who had received an ECG during pregnancy to those who had not. ECGs are readily available diagnostic tests in Ontario, with 4.7 million ECG diagnostic tests ordered in 2014 alone.²³ Less than a quarter of adult women in Ontario who have an annual health examination receive a routine ECG.¹⁸ The technical and professional components have fees of \$6.60 and \$4.45, respectively.²⁴ We identified ECGs performed during pregnancy through claims in the OHIP database using the specific procedure codes during the 40 weeks before delivery (G310 and G311). These codes have been used in prior studies to reliably identify physician claims for the procedure.¹⁸ We also used these codes to additionally identify earlier ECGs during the year prior to conception.

Additional characteristics

We defined additional baseline individual characteristics at the time of delivery by linking administrative healthcare databases. We obtained basic maternal demographic information including age, socioeconomic status, and home location using the Registered Persons Database, which is the registry of individuals eligible for provincial health insurance.^{25,26} We obtained hospitalization data from the Canadian Institute for Health Information Discharge Abstract Database.²⁷ We obtained data on outpatient clinic visits using the OHIP database.²⁸ We obtained further data about pregnancy and delivery for the maternal-newborn pair using the MOMBABY database. The date of conception was estimated by subtracting the gestational age at delivery from the date of delivery (index date).

Postpartum depression

The primary outcome was a physician diagnosis of postpartum depression (*ICD*-9 code 311) during the year after delivery determined through the validated OHIP database. This definition excluded psychotic disorders, substance abuse, or other psychiatric diagnoses. To minimize false-positive results, we required physician contact for a diagnosis of depression on two separate occasions in the first postpartum year. Using this stringent definition of two separate depression claims has a specificity of 94% and sensitivity of 61% (positive predictive value of 70%, negative predictive value of 92%).²⁹ Other studies have used similar approaches to ascertaining postpartum mental health diagnoses.^{30,31} We assumed that women without an ICD diagnosis code for depression did not have postpartum depression. The baseline incidence of postpartum depression among Canadian women with no history of mental illness is approximately 45 per 1000 when based on broader definitions of mental illness.³²

Additional diagnostic procedures

We used OHIP claims to identify five other non-invasive procedures in addition to an ECG ordered in pregnancy: echocardiogram, Holter monitor, chest X-ray, leg Doppler, and abdominal ultrasound.^{18,33–36} We also used OHIP claims to identify routine screening tests for pregnant women: urinalysis, glucose tolerance testing, and anemia. Information on prescription drugs was not available for women in our study due to the absence of universal drug coverage. Social services and psychology assessments were not available in the health services databases.

Statistical analysis

We used logistic regression to examine the incidence of postpartum depression within one year (binary outcome), comparing women who received an ECG during pregnancy to women who did not receive an ECG during pregnancy. Statistical testing examined associations before and after adjusting for measured baseline characteristics (Table 1) to assess for robustness in our risk estimates. Results based on proportional hazards analysis instead of logistic regression analysis yielded almost identical results and are shown as an additional test of robustness. Odds ratios were represented with 95% confidence intervals and a two-tailed type 1 error rate of 0.05 defined the threshold for statistical significance. All analyses were performed using SAS statistical software (v 9.4; SAS Institute, Cary, NC).

To test the robustness of our findings we explored how an association between prenatal ECG testing and postpartum depression might extend across a spectrum of severity. First, we performed a sensitivity analysis restricted to women with a term delivery, a normal hospital stay, no history of depression, heart disease, diabetes, no prior ECG, and no hospitalization in the previous year. Second, we performed an analysis where greater numbers of outpatient clinic visits for postpartum depression were required to meet criteria for a severe postpartum depression. Third, we examined the more stringent outcome of depression leading to a hospital admission (depression was listed as a contributing diagnosis but not necessarily the most responsible diagnosis). Fourth, we examined a broader outcome by including other mental health disorders in the year after delivery (codes 300–304, 305, 309, 311) (Appendix 1).

We also performed analyses to explore the frequency of depression in postpartum women who received additional other cardiovascular procedures: echocardiogram, Holter monitor, chest X-ray, leg Dopplers, and abdominal ultrasound. We assessed for dose response by examining the apparent impact of multiple rather than a single ECG test on the risk of postpartum depression. We examined the cascade of test ordering by evaluating the likelihood of women receiving subsequent diagnostic tests (e.g., echocardiogram) following an initial ECG. We performed a counterfactual specificity analysis examining the association between routine screening ordered in pregnancy (urinalysis, glucose, anemia) and postpartum depression.

Table I. Baseline characteristics.

		Electrocardiogram		
Variable	Value	Yes n = 157,352	No n = 3,080,866	
Demographic				
Age	<25 years	27,573 (17.5%)	531,894 (17.3%)	
	25-35 years	101,245 (64.3%)	2,099,229 (68.1%)	
	>35 years	28,534 (18.1%)	449,743 (14.6%)	
Socioeconomic status ^a	Higher	52,287 (33.2%)	1,134,465 (36.8%)	
	Middle	31,040 (19.7%)	621,683 (20.2%)	
	Lower	74,025 (47.0%)	1,324,718 (43.0%)	
Home location	Urban	144,238 (91.7%)	2,748,968 (89.2%)	
	Rural	13,114 (8.3%)	331,898 (10.8%)	
Medical #			,	
Antecedent ECG ^b	Yes	29,747 (18.9%)	185,965 (6.0%)	
Prior depression ^c	Yes	2488 (1.6%)	22,768 (0.7%)	
Heart disease ^d	Yes	26,279 (16.7%)	60,308 (2.0%)	
Diabetes ^e	Yes	12,828 (8.2%)	139,571 (4.5%)	
Clinic visits	Total <12	7,829 (5.0%)	420,963 (13.7%)	
	Total \ge 13	149,523 (95.0%)	2,659,903 (86.3%)	
Hospital admission	Yes	65,741 (41.8%)	1,077,165 (35.0%)	
Obstetric				
Parity	Primiparous	91,260 (58.0%)	1,984,016 (64.4%)	
	Multiparous	66,092 (42.0%)	1,096,850 (35.6%)	
Multiplicity	Singleton	152,175 (96.7%)	3,013,312 (97.8%)	
	Twins ^f	5,177 (3.3%)	67,554 (2.2%)	
Pregnancy duration	Preterm	13,260 (8.4%)	154,963 (5.0%)	
	At term	106,479 (67.7%)	1,833,801 (59.5%)	
	Post term	299 (0.2%)	7,036 (0.2%)	
	Missing	37,314 (23.7%)	1,085,066 (35.2%)	
Delivery route	Vaginal	100,364 (63.8%)	2,110,720 (68.5%)	
	Caesarean	48,456 (30.8%)	764,477 (24.8%)	
	Missing	8532 (5.4%)	205,669 (6.7%)	
Length of stay ^g	<3 days	93,554 (59.5%)	1,964,265 (63.8%)	
	>4 days	63,798 (40.5%)	1,116,601 (36.2%)	

 $^a\!Higher\!=\!top\ 2$ income quintiles; middle $=\!middle$ quintile; lower $=\!bottom\ two\ quintiles.$

#Denotes two or more visits in year before delivery.

^bDenotes electrocardiogram in year before conception.

^cCode 311.

^dCodes 390–429.

^eCode 250.

^fIncludes higher order multiples. ^gDays in hospital for delivery.

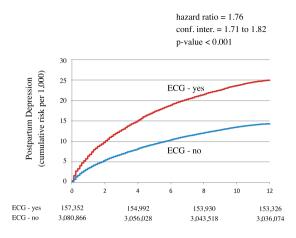
Results

Descriptive overview

In total, 3,238,218 women gave birth during the 25-year study period. Overall, 157,352 (nearly 1-in-20) received an ECG during prenatal care. Women spanned a wide range of socioeconomic status and the median age was 30 years (Table 1). The majority lived in a city and had no history of depression or heart disease. Most gave birth at term with a vaginal delivery and stayed in hospital less than four days. As expected, women who received an ECG relative to those who did not receive an ECG were slightly older, more likely to have a past history of heart disease, and more likely to have received an additional ECG in the year prior to pregnancy. The majority of women in both groups had more than a dozen prenatal care visits before delivery. The overall frequency of heart disease was 2.7%. Most ECG tests were in the first two trimesters (75%).

Subsequent depression

The women in our cohort accounted for 3,205,649.6 patient-years of subsequent follow-up (mean: 0.99 years). Women who received an ECG during prenatal care accounted for 3932 cases of subsequent depression over 154,597.6 patient-years of follow-up (mean: 0.98 years), equal to an incidence of 25 per 1000 annually. Women who did not receive an ECG during prenatal care accounted for 44,115 cases of subsequent depression over 3,051,052 patient-years of follow-up (mean: 0.99 years), equal to an incidence of 14 per 1000 annually. Together, receiving an ECG was associated with a 1.76 increased odds of subsequent depression (95% confidence interval)



Time (months following delivery)

Figure 1. Risk of postpartum depression. Cumulative incidence plots of absolute risk of postpartum depression. X-axis shows time following delivery spanning first year. Y-axis shows cumulative incidence of depression per 1000 individuals. Numerical counts show proportion of women with no depression at corresponding time. *P* value and hazard ratio based on Fine and Gray model. Results show substantial incidence of depression, particularly for women who received an ECG during prenatal care.

1.71–1.82, p < 0.001). About half of the postpartum cases of depression occurred after the first three months (Figure 1).

Other factors associated with depression

The risk of postpartum depression was also related to individual characteristics. Younger age, a rural home location, an ECG prior to conception, and a history of heart disease were each associated with increased odds of postpartum depression (Table 2). The strongest risk factor was a past history of depression. Additional predictors included an increased number of prenatal care visits, a prior hospital admission, being multiparous, a preterm delivery, and a longer hospital length of stay (perhaps each as indirect measures of comorbidity). Conversely, socioeconomic status was not a significant predictor after accounting for other factors. Adjustment for all measured women factors suggested that receiving an ECG test was associated with a 1.34 increased odds of postpartum depression (95% confidence interval 1.29–1.39, p < 0.001).

Subgroup analyses

We conducted a further check for confounding by replicating the analysis for women who had a term delivery; hospital stay \leq 3 days; no history of depression, heart disease, diabetes; no prior ECG; and no hospitalization in the past year. This resulted in a subgroup of 911,762 women of whom 34,054 (3.7%) received an ECG during prenatal care. Those who received an ECG accounted for 593 cases of postpartum depression over 33,666.8 patient-years of follow-up, equal to 18 per 1000 annually. Those who did not receive an ECG accounted for 10,238 cases of postpartum depression over 871,116.3 patient-years of follow-up, equal to 12 per 1000 annually. Together, this subgroup suggested receiving a prenatal ECG was associated with a 1.50 increased odds of postpartum depression (95% confidence interval 1.38–1.63, p < 0.001).

Secondary analyses

We conducted two further analyses to examine the validity of the association between receiving a prenatal ECG and risk of postpartum depression. The first analysis assessed a dose–response gradient by comparing women who received exactly two ECG tests to women who received exactly one ECG test and observed a further increase in depression (31 vs. 24 per 1000, p < 0.001). Analyses based on higher numbers of ECG tests yielded no further increases in risk (Figure 2). The second analysis assessed counterfactual specificity by comparing women who received a screening urinalysis (n = 2,969,795; 92%) to women who did not (n = 268,423; 8%) and observed only a slight increase in depression (15 vs. 14 per 1000, p < 0.001). Additional assessments based on glucose screening (15 vs. 14 per 1000, p < 0.001) screening yielded similar patterns.

Additional cardiovascular testing

We conducted further supplementary analyses to examine how receiving a prenatal ECG might lead to additional prenatal cardiovascular diagnostic procedures. Overall, we found a significant increase for each procedure comparing women who received a prenatal ECG test to women who did not receive a prenatal ECG test. The results were particularly striking for the likelihood of receiving a prenatal echocardiogram, Holter monitoring, and chest X-ray (Figure 3). In addition, the combination of prenatal ECG plus an additional prenatal cardiovascular diagnostic procedure compared to a prenatal ECG alone was associated with a further increased risk of postpartum depression (28 vs. 22 per 1000, p < 0.001).

Severity of outcome

We conducted additional analyses to examine the spectrum of postpartum depression by assessing indirect measures of disease severity. We found the association of ECG testing with postpartum depression extended to cases with many (>6) outpatient visits for depression treatment (5.5 vs. 2.7 per 1000, p < 0.001). Similarly, the association extended to cases with very many (>12) outpatient visits for depression treatment (1.4 vs. 0.7 per 1000, p < 0.001) and for cases with an acute care hospital admission for depression (4.2 vs. 1.9 per 1000, p < 0.001). The association also extended to a broader definition including other mental health diagnoses (159 vs. 103 per 1000, p < 0.001). In addition, the association extended to those who had a diagnosis of postpartum depression and died in the year following delivery, but the results were not statistically significant.

Discussion

We studied over three million pregnant women to assess whether an ECG test in pregnancy was associated with an increased risk of postpartum depression. The overall rate of ECG testing was about 1-in-20 and thereby far exceeded the baseline prevalence of heart disease in this demographic group.^{37,38} Compared to women who were not tested, women who received an ECG were at a one-third increased odds of postpartum depression after accounting for baseline predictors. The increased risk was further accentuated among pregnant women receiving additional cardiovascular procedures. In contrast, routine screening tests during pregnancy were not associated with a major increase in subsequent postpartum depression. Together, these findings suggest a possible link between physical and mental health.

One plausible explanation for the observed pattern of findings may be that somatic symptoms are a diagnostic clue to depression.^{39–42} For example, palpitations, dyspnea, and chest discomfort often prompt cardiovascular testing but can also be symptoms of a

Table 2. Predictors of depression in year following delivery.

Characteristic	Basic analysis ^a		Adjusted analysis ^b	
	Relative risk	Confidence interval	Relative risk	Confidence interval
Electrocardiogram test	1.76	1.71-1.82	1.34	1.29-1.39
Age < 25 years ^c	1.25	1.21–1.28	1.42	1.37–1.46
Age > 35 years ^c	0.91	0.88-0.93	1.02	0.99-1.05
Higher socioeconomic quintile ^d	0.96	0.93-0.98	0.99	0.96-1.01
Lower socioeconomic quintile ^d	1.04	1.01-1.06	1.00	0.97-1.02
Urban home location ^e	0.95	0.92-0.97	0.92	0.89-0.95
Antecdent ECG	1.72	1.67–1.77	1.34	1.30-1.38
Past depression	39.53	38.44-40.65	34.21	33.24–35.20
Past heart disease	1.42	1.36–1.49	1.05	1.00-1.11
Past diabetes	1.08	1.04-1.13	0.92	0.88–0.96
Clinic visits >13	1.73	1.67–1.78	1.59	1.54-1.65
Hospital admission	1.27	1.24-1.29	1.10	1.08-1.12
Multiparous ^f	1.28	1.26-1.31	1.10	1.08-1.13
Twins (or higher multiples)	1.43	1.36-1.51	0.97	0.92-1.03
Preterm pregnancy duration ^g	1.43	1.38–1.48	1.22	1.18–1.27
Post-term pregnancy duration ^g	1.01	0.84-1.20	1.07	0.90-1.29
Missing pregnancy duration ^g	0.60	0.59–0.1	0.64	0.63–0.66
Caesarean delivery route ^h	1.23	1.20-1.25	1.04	1.01-1.06
Missing data on delivery route ^h	1.18	1.14-1.22	1.34	1.29-1.39
Length of stay >4 days	1.30	1.27-1.32	1.24	1.21-1.27

Note: Estimates based on logistic regression model.

Analyses based on all women for all births.

^aNo adjustment for baseline differences.

^bAdjusted for all measures significant in univariable analysis.

^cReferent is age 25–35 years.

^dReferent is middle socioeconomic quintile.

^eReferent is rural home location.

^fReferent is primiparous.

^gReferent is at-term pregnancy. ^hReferent is vaginal.

mood disorder.^{43–45} Adults who have underlying structural heart disease may also be prone to subsequently developing a secondary depression.^{46–48} Physicians, for example, may anticipate the prospect of a mood disorder and decide to obtain an ECG before initiating formal psychiatric care. This means the association does not necessarily indicate causality and, instead, signals the presence of a baseline predilection to depression.

A second explanation could be that ordering a cardiovascular test in pregnancy creates a form of anchoring bias in clinicians who, in turn, become distracted from inquiring about mood symptoms or psychosocial stressors. Instead, prenatal care thereafter focuses on cardiovascular concerns and further diagnostic assessments may dominate the clinical agenda. This interpretation suggests that some women may be suffering in silence, leading to a delay in referral to appropriate mental health services.⁴⁹ This is an important reminder for clinicians to consider mental health diagnoses in pregnancy, particular when workup for cardiac conditions reveals no specific diagnosis.

A further explanation is that diagnostic tests during pregnancy are prone to false positive findings or ambiguous results (as occurs with screening mammography).^{50–52} Cardiovascular disease may be present in 0.2 to 4% of women during pregnancy.^{37,38} Therefore, ECG testing is sometimes indicated for appropriate prenatal care, including for women with preexisting heart disease, anesthetic planning, work-up for chest pain, and evaluation of palpitations. However, ECG irregularities are common and ectopic abnormalities appear in over half of pregnant women investigated for palpitations.^{53,54} Unfortunately, normal results are not always reassuring and cascades of diagnostic procedures might ensue, as observed in our study.^{18,55} Together, the implication is that fallible diagnostic test results could be distressing for individuals who expect a normal pregnancy and are not accustomed to extensive medical investigations.⁵⁶ Furthermore, the process of investigating a false positive test result can be stressful for women.

Our study also supports past research on the psychologically mediated effects of medical testing. Diagnostic tests are sometimes ordered to satisfy individual expectations or strengthen a therapeutic relationship.^{57–60} However, qualitative studies have sometimes shown the opposite.⁵⁹ Individuals can become distressed while waiting for results or from a cascade of ensuing tests.^{59,61–66} A systematic review and meta-analysis of studies suggested that diagnostic tests for symptoms with a low risk of serious illness do not reassure all individuals or reduce anxiety.^{65,67} Moreover, some women may suffer damaged self-identity by being labeled with an illness.^{68,69}

Our study has important limitations that merit emphasis. The study was not a randomized trial, and cannot establish how much of the association may reflect confounding from the indication for testing. To this end we have applied statistical analyses to adjust for variables such as preexisting heart disease, hypertension, and diabetes that might affect both the risk of postpartum depression and the

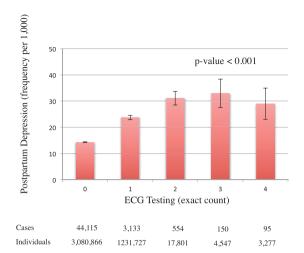


Figure 2. Dose response gradient. Frequency of postpartum depression stratified by number of ECG tests in prenatal interval. X-axis shows specific number ranging from 0 to 4 (final category also includes those with \geq 4). Y-axis shows cumulative risk of postpartum depression expressed per 1000 women. Vertical lines denote 95% confidence interval for each analysis. *P* value indicates test for trend. Numerical counts show total number of women in corresponding subgroup. Results show trend where greater number of ECG tests in prenatal care associated with greater frequency of postpartum depression.

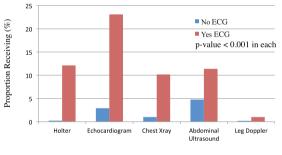




Figure 3. Additional diagnostic procedures. Absolute frequency of receiving additional prenatal diagnostic procedures among women who do or do not receive an ECG during prenatal care. X-axis shows specific diagnostic procedure of Holter monitor (codes G648 to G659, G682–G689), echocardiogram (G570, G571, G574, G575), chest X-ray (X090, X091, X092), abdominal ultrasound (J128, J135), and leg Doppler (J198). Y-axis shows proportion as percentage. Women who received an ECG during prenatal care indicated by red bar and remaining women indicated by blue bar. *P* value based on comparison of proportions in each procedure. Results show substantial frequency of additional prenatal diagnostic procedures for women who received an ECG during prenatal care.

likelihood of receiving an ECG.^{70–73} Regardless, the possibility of unmeasured confounders remain which might include tobacco smoking, alcohol consumption, substance misuse, domestic violence, and situational stress. This means an ECG could be a correlate or a marker of underlying comorbid illness, but not a contributor or cause of the increased risk of postpartum depression.

The lack of additional clinical information is important because our study detected only a modest association with ECG testing, far smaller than the association with a positive past history of depression.^{74,75} We also lacked information on other risk factors for postpartum depression including marital status,⁷⁶ intimate partner violence,^{70,77,78} breastfeeding challenges,^{79,80} psychological stress,^{81,82} and genetics.^{83,84} Furthermore, we did not have data on the concurrent management of depression including prescription medications, cognitive behavioral therapy, and prior electroconvulsive therapy. These uncertainties remain topics for future research.

Another further set of limitations in our research is that we may have failed to identify many women suffering from postpartum depression. The incidence of postpartum depression was measured using validated definitions that have imperfect specificity and sensitivity. The prevalence of postpartum depression is approximately 9% in North America, which is much higher than detected in our cohort.⁸⁵ This means our stringent definition may have missed many women who had a depressed mood, subclinical symptoms, or other psychiatric conditions. In addition, our study is based on one region in a high-income country that has free access to prenatal care and the findings may not apply elsewhere.

Conclusions

Our study identifies an association between prenatal ECG testing and postpartum depression, thereby highlighting a potential connection between physical symptoms and mental health in pregnancy. Physicians should be aware that cardiovascular symptoms may be a clinical clue to the presence of an underlying mood disorder in pregnant women that may become apparent postpartum.

Declaration of conflicting interests

The author(s) declared the following potential conflicts of interest with respect to the research, authorship, and/or publication of this article: JSZ has received fees for medicolegal opinions unrelated to the published work.

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Ethical approval

The study protocol was approved by the Research Ethics Board at Sunnybrook Health Sciences Centre, including a waiver for patient consent. Parts of this material are based on data and information compiled by CIHI.

Informed consent

Written consent was obtained from the patient(s) for their anonymized information to be published in this article.

Guarantor

DAR

Contributorship

JSZ and DAR contributed to the concept and design of the study. Data extraction and statistical analysis were performed by DT and DAR. JSZ and DAR drafted the manuscript. All authors performed revision of the manuscript.

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References

- Dietz PM, Williams SB, Callaghan WM, et al. Clinically identified maternal depression before, during, and after pregnancies ending in live births. *Am J Psychiatry* 2007; 164: 1515–20.
- Axfors C, Bränn E, Henriksson HE, et al. Cohort profile: the biology, affect, stress, imaging and cognition (BASIC) study on perinatal depression in a population-based Swedish cohort. *BMJ Open* 2019; 9: e031514.
- Bhati S and Richards K. A systematic review of the relationship between postpartum sleep disturbance and postpartum depression. J Obstet Gynecol Neonatal Nurs 2005; 44: 350–357.
- Committee on Obstetric Practice. The American College of Obstetricians and Gynecologists Committee Opinion no. 630. Screening for perinatal depression. *Obstet Gynecol* 2015; 125: 1268–1271.
- Yonkers KA, Vigod S and Ross LE. Diagnosis, pathophysiology, and management of mood disorders in pregnant and postpartum women. *Obstet Gynecol* 2011; 117: 961–977.
- Stewart DE. Clinical practice. Depression during pregnancy. N Engl J Med 2011; 365: 1605–1611.
- Stein A, Pearson RM, Goodman SH, et al. Effects of perinatal mental disorders on the fetus and child. *Lancet* 2014; 384: 1800–1819.
- Paulson JF, Dauber S and Leiferman JA. Individual and combined effects of postpartum depression in mothers and fathers on parenting behavior. *Pediatrics* 2006; 118: 659–668.
- Luca DL, Garlow N, Staatz C, et al. Societal Costs of Untreated Perinatal Mood and Anxiety Disorders in the United States. *Mathematica*, https://www.mathematica.org/our-publicationsand-findings/publications/societal-costs-of-untreated-perinatalmood-and-anxiety-disorders-in-the-united-states, (2019, accessed 6 November, 2020).
- Esscher A, Essén B, Innala E, et al. Suicides during pregnancy and 1 year postpartum in Sweden, 1980-2007. Br J Psychiatry 2016; 208: 462–469.
- Palladino CL, Singh V, Campbell J, et al. Homicide and suicide during the perinatal period: findings from the National Violent Death Reporting System. *Obstet Gynecol* 2011; 118: 1056–1063.
- Schiff MA and Grossman DC. Adverse perinatal outcomes and risk for postpartum suicide attempt in Washington state, 1987-2001. *Pediatrics* 2006; 118: e669–e675.
- ACOG Practice Bulletin No. 212: Pregnancy and heart disease. Obstet Gynecol 2019; 133: e320–e356.
- Adamson DL and Nelson-Piercy C. Managing palpitations and arrhythmias during pregnancy. *Heart* 2007; 93: 1630–1636.
- 15. Sunitha M, Chandrasekharappa S and Brid S. Electrocradiographic Q RS axis, Q wave and T-wave changes

in 2nd and 3rd trimester of normal pregnancy. J Clin Diagn Res 2014; 8: BC17–BC21.

- Hage B, Britton B, Daniels D, et al. Low cardiac vagal tone index by heart rate variability differentiates bipolar from major depression. *World J Biol Psychiatry* 2019; 20: 359–367.
- Hardavella G, Aamli-Gaagnat A, Frille A, et al. Top tips to deal with challenging situations: doctor-patient interactions. *Breathe* 2017; 13: 129–135.
- Bhatia RS, Bouck Z, Ivers NM, et al. Electrocardiograms in lowrisk patients undergoing an annual health examination. *JAMA Intern Med* 2017; 177: 1326–1333.
- Statistics Canada. Population and dwelling count highlight tables, 2006 census, http://www12.statcan.ca/census-recensement/2006/ dp-pd/hlt/97-550/Index.cfm?TPL = P3C&Page = INDX&LANG = Eng (2006, accessed November 6, 2020).
- Statistics Canada. Births, Canada, provinces and territories, 2006 to 2008, https://www150.statcan.gc.ca/n1/pub/84f0210x/ 2008000/t032-eng.htm (2012).
- Canadian Institute for Health Information. CIHI data quality study of emergency department visits for 2004-2005: *Executive Summary*. Ottawa, Ontario, 2008.
- 22. Fitzpatrick T, Wilton AS and Guttmann A. Development and validation of a simple algorithm to estimate common gestational age categories using standard administrative birth record data in Ontario, *Canada. J Obstet Gynaecol* 2020; 15(41): 1–5.
- Verma A, Crystal E, Dorian P, et al. Standards for provision of electrocardiography (ECG)-based diagnostic testing in Ontario. Toronto, 2017.
- Schedule of Benefits: Physician Services Under the Health Insurance Act, http://www.health.gov.on.ca/en/pro/programs/ ohip/sob/.
- 25. Wilkins R. Automated geographic coding based on the statistics Canada postal code conversion files, including postal codes to December 2003. Ottawa, Ontario, 2004.
- Iron K, Zagorski B, Sykora K, et al. Living and dying in Ontario: an opportunity for improved health information: ICES investigative report. Toronto, Ontario, Canada, 2008.
- 27. Juurlink D, Preyra C, Croxford R, et al. *Canadian Institute for Health Information Discharge Abstract Database: a validation study.* Toronto, 2006.
- Goel V, Williams J, Anderson G, et al. Summary of studies on the quality of health care administrative databases in Canada. In: *Patterns of health care in Ontario, the ICES practice atlas.* Ottawa, Ontario: Canadian Medical Association, 1996, pp. 339–345.
- Doktorchik C, Patten S, Eastwood C, et al. Validation of a case definition for depression in administrative data against primary chart data as a reference standard. *BMC Psychiatry* 2019; 19: 9.
- Steele LS, Glazier RH, Lin E, et al. Using administrative data to measure ambulatory mental health service provision in primary care. *Med Care* 2004; 42: 960–695.
- Vigod S, Sultana A, Fung K, et al. A population-based study of postpartum mental health service use by immigrant women in Ontario, Canada. *Can J Psychiatry* 2016; 61: 705–713.
- Brown HK, Wilton AS, Ray JG, et al. Chronic physical conditions and risk for perinatal mental illness: a population-based retrospective cohort study. *PLoS Med* 2019; 16: e1002864.
- Edwards JD, Kapral MK, Fang J, et al. Underutilization of ambulatory ECG monitoring after stroke and transient ischemic attack: missed opportunities for atrial fibrillation detection. *Stroke* 2016; 47: 1982–1989.
- Bouck Z, Mecredy G, Ivers NM, et al. Routine use of chest x-ray for low-risk patients undergoing a periodic health examination: a retrospective cohort study. *CMAJ Open* 2018; 6: E322–E329.

- 35. Thein H-H, Campitelli MA, Yeung LT, et al. Improved survival in patients with viral hepatitis-induced hepatocellular carcinoma undergoing recommended abdominal ultrasound surveillance in Ontario: a population-based retrospective cohort study. *PLoS One* 2015; 10: e0138907.
- Alotaibi GS, Wu C, Senthilselvan A, et al. The validity of ICD codes coupled with imaging procedure codes for identifying acute venous thromboembolism using administrative data. *Vasc Med* 2015; 20: 364–368.
- Elkayam U, Goland S, Pieper PG, et al. High-risk cardiac disease in pregnancy: part I. J Am Coll Cardiol 2016; 68: 396–410.
- 38. European Society of Gynecology (ESG), Association for European Paediatric Cardiology (AEPC), German Society for Gender Medicine (DGesGM), et al. ESC Guidelines on the management of cardiovascular diseases during pregnancy: the Task Force on the Management of Cardiovascular Diseases during Pregnancy of the European Society of Cardiology (ESC). Eur Heart J 2011; 32: 3147–3197.
- Silverstein B. Gender difference in the prevalence of clinical depression: the role played by depression associated with somatic symptoms. *Am J Psychiatry* 1999; 156: 480–482.
- Betrus PA, Elmore SK, Hamilton PA. Women and somatization: unrecognized depression. *Health Care Women Int* 1995; 16: 287–297.
- Kelly RH, Russo J and Katon W. Somatic complaints among pregnant women cared for in obstetrics: normal pregnancy or depressive and anxiety symptom amplification revisited? *Gen Hosp Psychiatry* 2001; 23: 107–13.
- Nylen KJ, Williamson JA, O'Hara MW, et al. Validity of somatic symptoms as indicators of depression in pregnancy. *Arch Womens Ment Health* 2013; 16: 203–10.
- Kapfhammer H-P. Somatic symptoms in depression. *Dialogues Clin Neurosci* 2006; 8: 227–239.
- Tylee A and Gandhi P. The importance of somatic symptoms in depression in primary care. *Prim Care Companion J Clin Psychiatry* 2005; 7: 167–176.
- 45. Tylee A, Gastpar M, Lépine JP, et al. DEPRES II (Depression Research in European Society II): a patient survey of the symptoms, disability and current management of depression in the community. DEPRES Steering Committee. Int Clin Psychopharmacol 1999; 14: 139–151.
- Terre L, Poston WSC, Foreyt J, et al. Do somatic complaints predict subsequent symptoms of depression? *Psychother Psychosom* 2003; 72: 261–267.
- Patten SB. Long-term medical conditions and major depression in a Canadian population study at waves 1 and 2. J Affect Disord 2001; 63: 35–41.
- Hotopf M, Mayou R, Wadsworth M, et al. Temporal relationships between physical symptoms and psychiatric disorder. Results from a national birth cohort. *Br J Psychiatry* 1998; 173: 255–261.
- Zauderer C. Postpartum depression: how childbirth educators can help break the silence. J Perinat Educ 2009; 18: 23–31.
- Molina Y, Beresford SAA and Thompson B. Psychological outcomes after a false positive mammogram: preliminary evidence for ethnic differences across time. *J Racial Ethn Heal Disparit* 2017; 4: 123–133.
- Salz T, Richman AR and Brewer NT. Meta-analyses of the effect of false-positive mammograms on generic and specific psychosocial outcomes. *Psychooncology* 2010; 19: 1026–1034.
- Bond M, Pavey T, Welch K, et al. Systematic review of the psychological consequences of false-positive screening mammograms. *Health Technol Assess* 2013; 17: 1–170, v–vi.
- Shotan A, Ostrzega E, Mehra A, et al. Incidence of arrhythmias in normal pregnancy and relation to palpitations, dizziness, and syncope. *Am J Cardiol* 1997; 79: 1061–1064.

- Gowda RM, Khan IA, Mehta NJ, et al. Cardiac arrhythmias in pregnancy: clinical and therapeutic considerations. *Int J Cardiol* 2003; 88: 129–133.
- 55. Bouck Z, Calzavara AJ, Ivers NM, et al. Association of low-value testing with subsequent health care use and clinical outcomes among low-risk primary care outpatients undergoing an annual health examination. JAMA Intern Med. Epub Ahead of print. 8 June 2020. DOI: 10.1001/jamainternmed.2020.1611.
- Zipursky JS and Redelmeier DA. Medical principles in obstetrical consults. *Am J Med* 2018; 131: 1405–1407.
- 57. Schneider J, Kaplan SH, Greenfield S, et al. Better physicianpatient relationships are associated with higher reported adherence to antiretroviral therapy in patients with HIV infection. *J Gen Intern Med* 2004; 19: 1096–1103.
- Mushlin AI, Kern LM, Paris M, et al. The value of diagnostic information to patients with chest pain suggestive of coronary artery disease. *Med Decis Making* 2005; 25: 149–57.
- Vis JY, van Zwieten MCB, Bossuyt PMM, et al. The influence of medical testing on patients' health: an overview from the gynecologists' perspective. *BMC Med Inform Decis Mak* 2013; 13: 117.
- Whiting P, Toerien M, de Salis I, et al. A review identifies and classifies reasons for ordering diagnostic tests. *J Clin Epidemiol* 2007; 60: 981–989.
- Sox HC, Margulies I and Sox CH. Psychologically mediated effects of diagnostic tests. *Ann Intern Med* 1981; 95: 680–685.
- 62. Cohn DM, Vansenne F, Kaptein AA, et al. The psychological impact of testing for thrombophilia: a systematic review. *J Thromb Haemost* 2008; 6: 1099–1104.
- 63. Giroldi E, Veldhuijzen W, Mannaerts A, et al. 'Doctor, please tell me it's nothing serious': an exploration of patients' worrying and reassuring cognitions using stimulated recall interviews. *BMC Fam Pract* 2014; 15: 73.
- Pineault P. Breast cancer screening: women's experiences of waiting for further testing. *Oncol Nurs Forum* 2007; 34: 847–853.
- 65. Kroenke K. Diagnostic testing and the illusory reassurance of normal results: comment on 'Reassurance after diagnostic testing with a low pretest probability of serious disease'. *JAMA Intern Med* 2013; 173: 416–417.
- Warner AS, Shah N, Morse A, et al. Patient and physician attitudes toward low-value diagnostic tests. *JAMA Intern Med* 2016; 176: 1219–1221.
- Rolfe A and Burton C. Reassurance after diagnostic testing with a low pretest probability of serious disease: systematic review and meta-analysis. *JAMA Intern Med* 2013; 173: 407–416.
- Haynes RB, Sackett DL, Taylor DW, et al. Increased absenteeism from work after detection and labeling of hypertensive patients. N Engl J Med 1978; 299: 741–744.
- Van Bulck L, Luyckx K, Goossens E, et al. Illness identity: capturing the influence of illness on the person's sense of self. *Eur J Cardiovasc Nurs* 2019; 18: 4–6.
- Gaillard A, Le Strat Y, Mandelbrot L, et al. Predictors of postpartum depression: prospective study of 264 women followed during pregnancy and postpartum. *Psychiatry Res* 2014; 215: 341–346.
- Katon W, Russo J and Gavin A. Predictors of postpartum depression. J Womens Health 2014; 23: 753–759.
- Kozhimannil KB, Pereira MA and Harlow BL. Association between diabetes and perinatal depression among low-income mothers. *JAMA* 2009; 301: 842–847.
- 73. Sundaram S, Harman JS and Cook RL. Maternal morbidities and postpartum depression: an analysis using the 2007 and 2008 pregnancy risk assessment monitoring system. *Womens Health Issues* 2014; 24: e381–e388.

- 74. O'Hara MW and McCabe JE. Postpartum depression: current status and future directions. *Annu Rev Clin Psychol* 2013; 9: 379–407.
- Milgrom J, Gemmill AW, Bilszta JL, et al. Antenatal risk factors for postnatal depression: a large prospective study. J Affect Disord 2008; 108: 147–157.
- O'Hara MW and Wisner KL. Perinatal mental illness: definition, description and aetiology. *Best Pract Res Clin Obstet Gynaecol* 2014; 28: 3–12.
- Paschetta E, Berrisford G, Coccia F, et al. Perinatal psychiatric disorders: an overview. Am J Obstet Gynecol 2014; 210: 501-509.e6.
- Kornfeld BD, Bair-Merritt MH, Frosch E, et al. Postpartum depression and intimate partner violence in urban mothers: cooccurrence and child healthcare utilization. *J Pediatr* 2012; 161: 348–53.e2.
- Chojenta CL, Lucke JC, Forder PM, et al. Maternal health factors as risks for postnatal depression: a prospective longitudinal study. *PLoS One* 2016; 11: e0147246.
- Dias CC and Figueiredo B. Breastfeeding and depression: a systematic review of the literature. J Affect Disord 2015; 171: 142–154.
- Beck CT. Predictors of postpartum depression: an update. Nurs Res 2001; 50: 275–285.
- Radesky JS, Zuckerman B, Silverstein M, et al. Inconsolable infant crying and maternal postpartum depressive symptoms. *Pediatrics* 2013; 131: e1857–e1864.
- Couto TCE, Brancaglion MYM, Alvim-Soares A, et al. Postpartum depression: a systematic review of the genetics involved. *World J Psych* 2015; 5: 103–111.

- Figueiredo FP, Parada AP, de Araujo LF, et al. The influence of genetic factors on peripartum depression: a systematic review. *J Affect Disord* 2015; 172: 265–273.
- Vesga-López O, Blanco C, Keyes K, et al. Psychiatric disorders in pregnant and postpartum women in the United States. *Arch Gen Psych* 2008; 65: 805–815.

Appendix I. Mental health ICD9 codes.

- 300: Neurotic disorders (e.g., anxiety, hysteria, obsessivecompulsive disorders)
- 301: Personality disorders (e.g., paranoid personality disorder, affective personality disorder)
- 302: Sexual deviations and disorders (e.g., pedophilia, exhibitionism)
- 303: Alcohol dependence syndrome
- 304: Drug dependence (e.g., cocaine dependence, cannabis dependence, opioid dependence)
- 305: Nondependent abuse of drugs (e.g., alcohol abuse, cannabis abuse, cocaine type abuse)
- 309: Adjustment reaction (e.g., brief depressive reaction, prolonged depression reaction)
- 311: Depressive disorder, not otherwise classified