Physiological Predictors of Postpartum Depression and Anxiety **ALLISON ERIKSSON** DOCTORAL CANDIDATE, INSTITUTION FOR WOMEN AND CHILDREN'S HEALTH WOMHER RESEARCH SCHOOL **UPPSALA UNIVERSITY** 

Background: Perinatal affective disorders

- DSM-5 definition of postpartum (peripartum) depression: Depressive episodes lasting longer than 2 weeks that occur during pregnancy or within 4 weeks after childbirth
- Estimated **10-20% of wome**n suffer from from postpartum affective disorders

• Associated with long-term health risks for both mother and child

## Background: Screening for PPD

- In a Swedish study, only 70% of women offered screening for depressive symptoms (Bränn et al., 2021)
- A systematic review of clinical recognition, treatment and treatment response in perinatal depression shows that only 30.8% of women with postpartum depression are identified in clinical settings; and very few, about 6-7%, receive adequate treatment (Cox et al., 2016)

## Difficulty recognizing when to seek help

Background: Barriers to care

 Stigmatization of mental illness: Main use of self-report measures could hinder the ability for clinicians to detect PPD.



• We can objectively measure the body's reaction to stress

 Pregnancy constitutes a significant mental and physical stressor

• Stress increases risk of affective disorders



## Physiological measures: Heart rate variability (HRV)

HRV is the measurement of beat-tobeat changes in heartrate using an ECG, and is thought to reflect emotional regulation and dysregulation and act as a proxy for the **autonomic nervous system** (ANS)

- $\rightarrow$  Parasympathetic (Vagus) : slows HR
- → Sympathetic : increases HR

Higher variability reflects the body's ability to respond and adapt to changes (regulation). Lower variability reflects poor adaptation to stressors (dysregulation).

#### **NERVOUS SYSTEM**





# HRV and Depression/Anxiety



- HRV indices were found to exhibit screening sensitivity of 80% for Major Depressive Disorder compared to the subjective patient-reported screening method in non-pregnant sample (Sun et al., 2016)
- Previous studies have shown lower HRV to be associated with symptoms of affective disorders in non-pregnant samples (Choi et al., 2019) and vice versa (Jandackova et al., 2016)
- HRV during pregnancy is altered in association with past or current anxiety disorders as well as with greater trait anxiety, or greater exposure to past traumatic events (Kimmel et al., 2021)

However, the **predicitive** value of HRV for affective disorders has not been studied in pregnant women.

# Project Purpose

• Early identification is crucial.

 This study investigates if an objective measure (i.e. heart rate variability (HRV)) during pregnancy is related to later development of postpartum affective disorders Data from Uppsala University's **BASIC (Biology, Affect, Stress, Imaging and Cognition)** cohort

- 115 women
- Predictor: HRV measurements before and after a mild stressor at gestational week 38
- Outcome: Depressive and anxiety symptoms from 6 weeks postpartum measured using web surveys, including the Edinburgh Postnatal Depression Scale (EPDS)

**Purpose**: To test the predictive power of HRV in **late pregnancy** for PPD and anxiety, establishing important design and analysis

parameters for subsequent studies.



## HRV Indices

Time-Domain

- SDNN (standard deviation of normal-normal intervals)
  - RMSSD (Root mean square of successive differences)

#### Frequency-Domain

- Low frequency (LF: 0.04 ~ 0.15 Hz)
  - High frequency (HF: 0.15 ~ 0.4 Hz)

- Total power
- ●/● LF/HF ratio

- Reflects sympathetic activity
- Reflects parasympathetic activity



### Depression:

Edinburgh Postnatal Depression Scale (EPDS)(pregnancy w. 38, pp w. 6)

Anxiety:

- Spielberger State Trait Anxiety Inventory (STAI) (preg w.38)
- Beck Anxiety Inventory (pp w.6)



## **Statistics**



Preliminary analyses: Chi-square and independent samples t-tests

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Planned: Linear and logistic regression modeling

### Preliminary Results: Background variables

| Background<br>Variables       | Healthy<br>n (%) |             |      |
|-------------------------------|------------------|-------------|------|
| Place of birth                |                  |             |      |
| Scandinavia                   | 79 (95.2)        | 25 (92.6)   |      |
| Other                         | 4 (4.8)          | 2 (7.4)     | .607 |
| Maternal age at time of birth |                  |             |      |
| 21-29                         | 29 (33.3)        | 10 (37)     |      |
| 30-34                         | 33 (37.9)        | 14 (51.9)   |      |
| 35-42                         | 25 (28.7)        | 3 (11.1)    | .161 |
| Education                     |                  |             |      |
| University                    | 65 (76.5)        | 23 (85.2)   |      |
| Less                          | 20 (23.5)        | 4 (14.8)    | .336 |
| BMI*                          | 82               | 26          |      |
| Mean (sd)                     | 28.16 (3.9)      | 28.65 (4.4) | .610 |
| Illness combined*             |                  |             |      |
| No                            | 62 (81.6)        | 20 (83.3)   |      |
| Yes                           | 14 (18.4)        | 4 (16.7)    | .845 |
| Previous depression           |                  |             |      |
| Depression history            | 57 (67.9)        | 23 (85.2)   |      |
| No depression history         | 27 (32.1)        | 4 (14.8)    | .081 |

### Preliminary Results: Pregnancy variables

| Pregnancy<br>Variables      | Healthy<br>n (%) | Depressed<br>n (%) | p-value |
|-----------------------------|------------------|--------------------|---------|
| Sleeping habits week 32     |                  |                    |         |
| Less than 6 hours           | 6 (6.9)          | 4 (14.3)           |         |
| 6 -8 hours                  | 63 (72.4)        | 14 (50)            |         |
| More than 8 hours           | 18 (20.7)        | 10 (35.7)          | .087    |
| Fear of delivery week 32    |                  |                    | -       |
| No                          | 84 (96.6)        | 23 (82.1)          |         |
| Yes                         | 3 (3.4)          | 5 (17.9)           | .009*   |
| SSRI use at week 32         |                  |                    | -       |
| No                          | 73 (98.4)        | 23 (88.5)          |         |
| Yes                         | 4 (5.2)          | 3 (11.5)           | .266    |
| Depressive symptoms week 38 |                  |                    |         |
| 0 - 11                      | 66 (85.7)        | 12 (50)            |         |
| 12 - 30                     | 11 (14.3)        | 12 (50)            | <.001*  |
| State anxiety week 38       |                  |                    |         |
| STAI-S 6 - 11               | 75 (89.3)        | 15 (57.5)          |         |
| STAI-S 12 -24               | 9 (10.7)         | 11 (42.3)          | <.001*  |
| Trait anxiety week 38       |                  |                    |         |
| STAI-T 20 -39               | 67 (79.8)        | 8 (32)             |         |
| STAI-T 40 -80               | 17 (20.2)        | 17 (68)            | <.001*  |

### Preliminary Results: Postpartum variables

| Postpartum<br>Variables | Healthy<br>n (%) | Depressed<br>n (%) | p-value |  |
|-------------------------|------------------|--------------------|---------|--|
| Baby to neonatal unit   |                  |                    |         |  |
| No                      | 73 (94.8)        | 22 (88)            |         |  |
| Yes                     | 4 (5.2)          | 3 (12)             | .242    |  |
| Problems after delivery |                  |                    |         |  |
| (mother)*               |                  |                    |         |  |
| No                      | 54 (62.1)        | 9 (32.1)           |         |  |
| Yes                     | 33 (37.9)        | 19 (67.9)          | .006*   |  |
| Breastfeeding           |                  |                    |         |  |
| No                      | 4 (4.6)          | 4 (14.3)           |         |  |
| Yes                     | 83 (95.4)        | 24 (87.5)          | .080    |  |

|   |  | HRV Category                          | Type of scale | Healthy |                    | Depressed/Anxious |                                      | p -value        |
|---|--|---------------------------------------|---------------|---------|--------------------|-------------------|--------------------------------------|-----------------|
|   |  | int, category                         | ••            | n       | M <i>(SD)</i>      | n                 | M (SD)                               | • 0 00000000000 |
|   |  | RMSSD 1                               | Depression    | 87      | 32.7 (27)          | 28                | 23.0 (11)                            | .068            |
|   |  |                                       | Anxiety       | 95      | 29.2 (15.8)        | 14                | 36.1 <i>(</i> 55 <i>.</i> 3 <i>)</i> | .328            |
|   |  | RMSSD 2                               | Depression    | 86      | 33.7 (19.2)        | 26                | 28.1 <i>(13.9)</i>                   | .171            |
|   |  |                                       | Anxiety       | 94      | 32.8 (17.6)        | 12                | 24.8 (14.1)                          | .135            |
|   |  | SDNN 1                                | Depression    | 87      | 56.2 (22)          | 28                | 47.2 (21)                            | .060            |
|   |  |                                       | Anxiety       | 95      | 53.9 (20.1)        | 14                | 51.8 (32.6)                          | .740            |
|   |  | SDNN 2                                | Depression    | 86      | 55.7 (21)          | 26                | 46.6 (17)                            | .050*           |
|   |  |                                       | Anxiety       | 94      | 54.3 (20.6)        | 12                | 41.5 (13.4)                          | .039*           |
| <u>Preliminary</u>                      |  | HF power 1                            | Depression    | 87      | 5.93 (1.2)         | 28                | 5.25 (1.3)                           | .013*           |
| -                                       |  |                                       | Anxiety       | 95      | 5.8 (1.6)          | 14                | 5.44 (1.6)                           | .284            |
| Results:                                |  | HF power 2                            | Depression    | 86      | 6.09 (1.1)         | 26                | 5.64 (1.1)                           | .071            |
| <u>Nesuits.</u>                         |  | III power 2                           | Anxiety       | 94      | 6.04 (1.0)         | 12                | 5.45 (1.3)                           | .071            |
| HRV                                     |  | LF power 1                            | Depression    | 87      | 6.03 (.8)          | 28                | 5.57 (1.0)                           | .018*           |
|   |  | · · · · · · · · · · · · · · · · · · · | Anxiety       | 95      | 5.94 (.80)         | 14                | 5.75 (1.1)                           | .455            |
| indices                                 |  | LF power 2                            | Depression    | 86      | 6.1 (.74)          | 26                | 5.9 (.92)                            | .377            |
| <u>IIIUICES</u>                         |  | LI power 2                            | Anxiety       | 94      | 6.07 (.76)         | 12                | 5.9 (.82)                            | .463            |
|   |  | Total power 1                         | Depression    | 87      | 7.89 (. <i>7</i> ) | 28                | 7.53 (.8)                            | .023*           |
|   |  |                                       | Anxiety       | 95      | 7.83 (.7)          | 14                | 7.57 (.8)                            | .212            |
|   | _  | T- 4-1 0                              | Depression    | 86      | 7.88 (7)           | 26                | 7.53 (0.7)                           | .051            |
|   |  | • Total power 2                       | Anxiety       | 94      | 7.83 (.7)          | 12                | 7.35 (.65)                           | .034*           |
|   |  | LF/HF ratio 1                         | Depression    | 87      | 1.6 (1.6)          | 28                | 1.7 (1.5)                            | .625            |
| = Reflects sympathetic activity         |  | LIT/IIT TAUD I                        | Anxiety       | 95      | 1.62 (1.6)         | 14                | 1.88 (1.7)                           | .569            |
| Deflects as a second state stick stick. | N/   | I E/IIE motic 2                       | Depression    | 86      | 1.45 (1.3)         | 26                | 1.97 (2.2)                           | .144            |
|   | eflects parasympathetic activity LF/HF ratio 2 |                                       | Anxiety       | 94      | 1.52 (1.6)         | 12                | 2.09 (1.7)                           | .252            |

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# <u>Conclusions</u>

 Preliminary data show a relationship between HRV indices measured at pregnancy week 38 and later development of depressive symptoms during the postpartum period.

Moving forward, we would like to use this data to:

- create a multivariate model adjusting also for pregnancy symptoms, and using stricter criteria for depressive outcomes
- test the predictive power of HRV together with self reports

#### **Clinical Relevance:**

 Can HRV improve a model for predicition of affective disorders in the postpartum period?

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