Data-Driven Insights towards Risk Assessment of Postpartum Depression

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Abstract:

Postpartum depression is defined as depressive episodes that occur during pregnancy or within 12 months of parturition. The goal of this study is the exploration of the birth features and maternal traits which affect the risk of postpartum depression for mothers with preterm neonates. We analysed data from 144 women (63 mothers of term and 81 mothers of preterm infants) who completed the Edinburgh Postnatal Depression Scale (EPDS) in the postpartum period. We used three feature selection algorithms: *ReliefF, Random Forests (RF) variable importance*, and *Boruta*, in order to select the most predictive feature subsets, which were subsequently mapped onto the binarized EPDS total score (a threshold of 10 was used to binarize the EPDS total scores) using RF. We found that positive affectivity (r_s =-0.467, p<0.001), and the Apgar score at 5 minutes (r_s =-0.430, p<0.001) are the most statistically strongly associated features with the risk of postpartum depression. We used 10-fold cross-validation with 100 iterations and report out-of-sample balanced accuracy (median±IQR): 75.0±16.7, sensitivity: 66.7±16.7, specificity: 100±16.7, and F1 score: 0.8±0.2. Collectively, these findings highlight the potential of using a data-driven process to automate risk prediction using standard clinical characteristics and motivate the deployment of the developed tool using larger-scale datasets.

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1 INTRODUCTION

Non-psychotic mental disorders are common during pregnancy and the postpartum period, and have a negative impact on the mother and her offspring (Howard *et al.*, 2014). Postpartum depression is defined by the American Psychiatric Association's Diagnostic and Statistical Manual, Fifth Edition (DSM-5) as a major depressive episode that occurs either during pregnancy or within 4 weeks of parturition (APA, 2013). In practice, it is operationalised as depressive episodes occurring within 6 to 12 months of delivery (Committee on Obstetric Practice, 2015). Postnatal non-psychotic depression affects approximately 3-6% of women during pregnancy or in the postnatal period (APA, 2013).

Clinical manifestations include despondency, emotional lability, tearfulness, feelings of guilt, anhedonia, low energy, loss of appetite, poor sleep, poor concentration and memory, fatigue, irritability, as well as feelings of being incapable of looking after the offspring (Stewart, 2005). It is also common for some women to report suicidal ideation (Martini *et al.*, 2019). Many studies have shown that mothers' perinatal depressive symptoms can negatively affect the social, emotional, and cognitive developmental outcomes of their offspring (Letourneau *et al.*, 2019).

Risk factors for postpartum depression include previous history of mental health disorders such as depression and anxiety, negative feelings towards the current birth, low self-esteem regarding the parental role, and positive history of sexual abuse (Ghaedrahmati et al., 2017). Risky pregnancy, emergency caesarean section, complications and long hospitalization also seem to have a negative impact on mothers' mental health, whereas breastfeeding is associated with lower risk of depression. Furthermore, studies have shown a positive correlation between young age during disorders, pregnancy, metabolism reduced serotonin, oxytocin, oestrogen levels and postnatal depressed mood (Ghaedrahmati et al., 2017). Finally, lack of emotional and financial support, unemployment and low socioeconomic status, sexual and domestic violence during the prenatal period, sleeping disorders, as well as habits such as smoking, low physical activity and unhealthy diet can potentially increase the risk of postpartum mood disorders (Ghaedrahmati et al., 2017).

Early detection of postpartum depression is alarmingly low (Moraes *et al.*, 2017). One standard widely used clinical metric which is a validated

screening tool for postpartum depression is the Edinburgh Postnatal Depression Scale (EPDS) (Cox, Holden and Sagovsky, 1987). It consists of ten items which assess symptoms of depression such as anxiety, low energy, sleep disorders, and suicidal thoughts. The scale shows how the mother has felt over the preceding week. Total scores of ten or more indicate possible depression of varying severity. The EPDS may be used within eight weeks postpartum and it can also be used as a depression screening tool during pregnancy (Cox, Holden and Sagovsky, 1987). According to a systematic review of validation studies of the EPDS conducted by Gibson et al. 2009, when using a cut-off point of 10, the sensitivity of the EPDS for detecting postpartum depression ranged from 59 to 100%, and the specificity ranged from 44 to 97% (Gibson et al., 2009).

The aim of this study is to explore the birth features and maternal traits that affect the risk of postpartum depression for mothers and build a statistical learning model in order to predict the risk of postpartum depression for mothers who have given birth to preterm neonates.

2 DATA

The study used data that comes from a longitudinal study conducted at the Royal Infirmary of Edinburgh which investigates the impact of preterm birth on long-term outcomes (https://theirworld.org). A total of 144 women completed the EPDS during the first month following childbirth, of whom 63 had term born neonates, and 81 had preterm neonates. We analysed data comprising demographic characteristics, characteristics (i.e. gestational age, infants' birthweight, occipitofrontal circumference, Apgar score), morbidities, and complications that occur during the neonatal period, socioeconomic status, and data derived from questionnaires which assess characteristics of temperament (Adult Temperament Questionnaire, short form), intelligence (National Adult Reading Test), and risk of postpartum depression (Edinburgh Postnatal Depression Scale). Socioeconomic status of the family was determined using the Scottish Index of Multiple Deprivation 2016. Table 1 presents the demographic characteristics of the mothers.

| | Mothers with | Mothers with |
|---------------------|------------------|-------------------|
| Characteristic | term born | preterm |
| Characteristic | neonates | neonates |
| | (n=63) | (n=81) |
| Age (years) | 34 ± 7 | 31 ± 7 |
| Ethnicity | | |
| Any White | 54 (91) | 73 (93) |
| Asian | 2 (4) | 4 (5) |
| Black | 0 (0) | 1(1) |
| Mixed | 2 (3) | 1(1) |
| Other | 1 (2) | 0 (0) |
| Education | | |
| None | 0 (0) | 2 (3) |
| 1-4 GCSE | 1 (2) | 4 (6) |
| \geq 5 GCSE | 2 (3) | 5 (7) |
| A Levels/Highers | 2 (3) | 9 (12) |
| College | 4 (7) | 17 (24) |
| University | 27 (47) | 22 (31) |
| Postgraduate degree | 22 (38) | 13 (18) |
| SIMD Quintile | | |
| 1 | 3 (5) | 10 (13) |
| 2 | 8 (13) | 22 (28) |
| 3 | 11 (17) | 12 (15) |
| 4 | 17 (27) | 18 (23) |
| 5 | 24 (38) | 16 (21) |
| Full IQ | 122.74 ± 4.96 | 121.10 ± 7.43 |
| BMI | 24.7 ± 6.6 | 26.2 ± 7.28 |
| Gestation (weeks) | 39.56 ± 1.42 | 29.84 ± 2.72 |

Table 1: Demographic characteristics of the mothers who completed the EPDS in the postpartum period.

Variables are presented in the form median ± IQR or number (%). Education refers to the highest educational level attained. GCSE stands for General Certificate of Secondary Education. SIMD stands for Scottish Index of Multiple Deprivation. IQ stands for Intelligence Quotient. BMI stands for Body Mass Index.

3 METHODS

We used the standard clinical cut-off of 10 (Cox, Holden and Sagovsky, 1987) to binarize the EPDS total scores; scores lower than 10 were classified as "low risk for postpartum depression", and scores equal to or higher than 10 were classified as "high risk for postpartum depression". We aimed to build a statistical model in order to estimate whether scores would be above or below the clinical threshold (i.e. this is a binary classification task).

3.1 Preliminary Statistical Analysis

We used the *Mann – Whitney rank sum test* (Mann and Whitney, 1947) to test whether the difference

between the medians of the EPDS total score of mothers with term born neonates and mothers with preterm neonates is zero. The level of significance associated with the null hypothesis was set at $\alpha = 0.05$.

In order to quantify the strength of the association between two variables, we computed the *Spearman's rank correlation coefficient* which is effective in quantifying general monotonic relationships (Schober, Boer and Schwarte, 2018). There is no universal guideline to determine when a bivariate relationship is statistically strong; it depends on the application (Cohen *et al.*, 2002). In this study, we consider that an absolute value of a correlation coefficient > 0.3 corresponds to a statistically strong association, in accordance to similar studies in clinical contexts (Meyer *et al.*, 2001; Tsanas, Little and McSharry, 2013).

3.2 Feature Selection

A problem that often arises when we analyse high dimensional data is the curse of dimensionality (Bellman, 1966); given that there is only a finite number of available samples, as the dimensions of the feature space increase, it is difficult to adequately populate the feature space, with detrimental effects in the performance of the learners (Hastie, Tibshirani and Friedman, 2009). The number of data samples required, often grows exponentially with the number of features. Prediction performance can often be improved by reducing the number of dimensions in the feature space, a process that is called dimensionality reduction. Dimensionality reduction techniques can be divided into two main categories: feature transformation and feature selection (Guyon and Elisseeff, 2003).

In this study, we have compared three efficient feature selection algorithms: (a) *ReliefF* (Kononenko, 1994), (b) *Random Forests (RF) variable importance* (Hastie, Tibshirani and Friedman, 2009; James *et al.*, 2013), and (c) *Boruta* (Kursa and Rudnicki, 2010).

ReliefF is a *feature-weighting algorithm*; it assigns a 'weight' value to all features of a dataset based on how well their values distinguish between the data samples that are near to each other and thus, how useful they are in predicting the response variable. There are various feature evaluation measures that ReliefF can use. In this study, we have explored ReliefF *expRank* where k nearest instances have weight exponentially decreasing with increasing rank.

When using RF (Breiman, 2001), it is possible to measure the importance of each feature in predicting

the response variable. At each split in each tree, the improvement in the split – criterion is the importance measure attributed to the splitting variable and is accumulated over all the trees in the forest for each feature. In the context of classification models, we can measure the total amount that the Gini index is decreased by splits over a given feature, averaged over all trees. A large value indicates an important feature (Hastie, Tibshirani and Friedman, 2009; James *et al.*, 2013).

The Boruta algorithm is a wrapper feature selection technique built around the random forest learner that measures the importance of each feature by dividing the average loss of accuracy among all trees by the standard deviation of the accuracy loss (Kursa and Rudnicki, 2010). In other words, it uses Z score as the importance measure.

In this study, the feature subsets were selected using a Cross-Validation (CV) approach, using only the training dataset in each CV iteration. We repeated the CV process a total of 100 times. The feature selection algorithms described above aim at ranking the features of a given dataset based on their contribution towards prediction of the response variable. In order to select the final feature subset for each feature selection algorithm, we followed the process described by Tsanas et al. in 2012 (Tsanas et al., 2012). In a nutshell, for a given feature selection algorithm, when using CV, at the end of each iteration, we obtain a vector of the ordered sequence of the indices of the features, where the first feature is considered to be the most important one, and the last feature corresponds to the least important one. We store these vectors in a matrix of $n \ge p$ size, where ncorresponds to the number of iterations and p corresponds to the number of features of a given dataset. This way, in each of the rows of the matrix we have stored the feature subset selected at the end of each iteration. Subsequently, we need to identify the feature index which has most frequently been ranked as first across all iterations, then we need to identify which feature appears most frequently as second or third and so on. In case a feature index has already been included in the final subset and is later found again as most frequent, we need to select the second most frequent and so on. Ties are resolved by including the lowest index number. In the end, we obtain a vector which consists of feature indices ordered from the most important to the least important one, for a given feature selection algorithm. These features are then used to train the learner one by one. We choose to keep the most parsimonious model that gives the best overall results in terms of accuracy, sensitivity, specificity, and F1 score.

3.3 Mapping Selected Features to the Response

Our ultimate goal is to estimate the function f which relates **X** and **y**, that is $f(\mathbf{X}) = \mathbf{y}$, where **y** is the response variable and **X** is a matrix comprising p features $X_1, X_2, X_3, ..., X_p$. That is, we need a binary classifier that will use the selected features to predict the risk of postpartum depression.

Here, we used RF (Breiman, 2001) which is a widely-used statistical machine learning algorithm. RF involve fitting decision trees on bootstrapped samples of the original training data and then combining all individual trees (we used 500 trees) to create a single powerful predictive model. When building these decision trees, each time a split in a tree is considered, a random sample of m features is chosen as split candidates from the full set of p features. The split is allowed to use only one of those m predictors. Following the suggestion of Breiman (Breiman, 2001), we set m to be equal to the square root of the number of features in the training data.

3.4 Classifier Validation

We validated our model using CV, which provides an estimate of the performance of the model on new "unseen" data, provided that the new data comes from the same joint distribution as the data that was used to train the model. In this study, we used 10-fold cross-validation and repeated the process 10 times, resulting in 100 iterations.

4 RESULTS

4.1 Preliminary Statistical Analysis

Regarding mothers who had term born neonates (n = 63), the total scores of the EPDS ranged from 0 to 15, with a median score of 6. Among them, 16 mothers had a total score of 10 or above, indicating high risk of depression, while the rest scored lower than 10. With regards to the mothers who had preterm neonates (n = 81), the total scores of the EPDS ranged from 0 to 23, with a median score of 7. Among them, 27 mothers had a total score of 10 or above, indicating high risk of depression, while the rest of the mothers scored lower than 10.

Figure 1 shows a histogram of the total scores of the EPDS for mothers with term born neonates and mothers with preterm neonates. The dashed lines represent the median scores. The p-value of the Mann - Whitney rank sum test is 0.459. At the 0.05 significance level, there is not enough evidence to reject the null hypothesis that there is no statistically significant difference in the medians of the EPDS total score between mothers who had a term born neonate and those who gave birth to preterm neonates.



Figure 1: EPDS total score histogram plot. The dashed line represents the median scores.

Table 2: Correlation analysis for the EPDS total score of the mothers who had preterm neonates (n = 81).

| Feature | Statistical association (Spearman correlation coefficient) | Statistical significance (p-value) |
|--------------------------------|--|--|
| Positive affect | -0.467 | < 0.001 |
| Apgar 5min | -0.430 | < 0.001 |
| Fear | 0.399 | < 0.01 |
| Extraversion | -0.347 | < 0.01 |
| Attentional control | -0.339 | < 0.01 |
| Days of intubation | 0.330 | < 0.01 |
| Days of supplemental O2 | 0.323 | < 0.01 |
| Gestation | -0.318 | < 0.01 |
| Previous live births | 0.309 | < 0.01 |
| Neutral perceptual sensitivity | -0.307 | <0.05 |

Only the ten most strongly associated features with the EPDS total score are presented in the Table for brevity. Positive affect, fear, extraversion, attentional control, and neutral perceptual sensitivity are characteristics of temperament derived from the Adult Temperament Questionnaire, short form, and assessed during the first month following delivery.

Table 2 presents the ten features (out of 102) most strongly associated with the total score of the EPDS for mothers with preterm infants. The features are sorted according to the absolute value of the correlation coefficient. Positive affect, fear, extraversion, attentional control, previous live births, and neutral perceptual sensitivity are all maternal features. Apgar score at 5 minutes, total number of days of intubation, total number of days of supplemental oxygen, and gestation refer to infant features. It is interesting to note that all features exhibit statistically significant (p<0.05) correlation. These findings give some initial confidence that the binary classification task of this study may lead to accurate results.

4.2 Classification Stage: Mapping Features to the Thresholded EPDS Total Score for Mothers with Preterm Neonates

Figures 2, 3 and 4 illustrate the out of sample performance of the RF as a function of the number of features selected by the different feature selection algorithms. We found that the feature size giving the best results in terms of accuracy, sensitivity, specificity, and F1 score is 13 using ReliefF expRank. Therefore, our subsequent results use the first 13 features selected by each feature selection algorithm. Table 4 presents the selected feature subsets for each feature selection algorithm. We remark that reducing



Figure 2: Comparison of out of sample median balanced accuracy results with IQR of the random forests classifier using the features selected by each of the five feature selection algorithms. These results are computed using 10-fold CV with 100 iterations. For clarity, only the first 30 steps are presented.

the original 102 dimensions of the feature space leads to an improvement in out-of-sample performance accuracy with RF. Table 3 shows a confusion matrix of the performance of the RF when mapping the features selected by ReliefF expRank (i.e. previous live births, bronchopulmonary dysplasia, and extraversion) to the response.



Figure 3: Comparison of out of sample median sensitivity results with IQR of the random forests classifier using the features selected by each of the five feature selection algorithms. These results are computed using 10-fold CV with 100 iterations. For clarity, only the first 30 steps are presented.



Figure 4: Comparison of out of sample median specificity results with IQR of the random forests classifier using the features selected by each of the five feature selection algorithms. These results are computed using 10-fold CV with 100 iterations. For clarity, only the first 30 steps are presented.

Table 3: Confusion matrix summarizing the out-of-sample findings using 10-fold CV.

| | | Reference | |
|------------|-------------------------|-------------------------|------------------------|
| | | High risk of depression | Low risk of depression |
| Prediction | High risk of depression | 165 | 47 |
| | Low risk of depression | 105 | 493 |

Total number of entries is 810, which is derived from 81 samples x 10 iterations.

Table 4: Selected feature subsets and out-of-sample classification performance.

| | ReliefF | RF variable | Boruta |
|-------------|--------------------------------------|--|--|
| | expRanK | importance | |
| | Previous live births | Extraversion | Extraversion |
| | BPD | Positive affect | Positive affect |
| | Extraversion | Fear | Attentional control |
| | Maternal anxiety | Gestation | Fear |
| | Days of intubation | Attentional control | Gestation |
| | Apgar score | Inhibitory | Apgar score at |
| | at 5 min | control | 5 min |
| | Days of | Effortful | Inhibitory |
| | supplemental | control | control |
| LOGY | O2 | | |
| | Effortful control | Apgar score at 5 min | Effortful control |
| | Inhibitory | Affective | Affective |
| | control | perceptual sensitivity | perceptual sensitivity |
| | Positive affect | Days of supple- mental O ₂ | Days of supple- mental O ₂ |
| | Feeding at discharge from NICU | Previous live births | Days of intubation |
| | ROP | Negative affect | Negative affect |
| | Maternal asthma | Total days in the NICU | Total days in the NICU |
| Accuracy | 75.0±16.7 | 83.3±18.3 | 75.0±23.3 |
| Specificity | 100±16.7 | 100±16.7 | 100±20.0 |
| Sensitivity | 66.7±16.7 | 66.7±50.0 | 66.7±66.7 |
| F1 score | 0.8 ± 0.2 | 0.8±0.2 | 0.8±0.2 |

The selected features from each feature selection algorithm are sorted in descending order from the most important to the least important one. The last row presents the performance of the random forests classifier for each feature subset. The results are given in the form median±IQR. BPD stands for Bronchopulmonary Dysplasia. NICU stands for Newborn Intensive Care Unit. ROP stands for Retinopathy of Prematurity.

5 DISCUSSION

In this study, our goal has been to build a parsimonious model to determine the features that could potentially predict the risk of postpartum depression for mothers with preterm infants. To this end, we compared three different feature selection algorithms: (a) ReliefF, (b) Boruta, and (c) RF variable importance in order to derive a small feature subset which would be more informative than the original feature set. Subsequently, in order to map the selected features to the response we used RF.

We analysed data from 144 women who completed the EPDS during the postpartum period, of whom 63 have had term born neonates, and 81 have had preterm neonates. For mothers who gave birth to preterm neonates, correlation analysis revealed statistically strong correlations. At this point, we need to emphasize that interpretation of the results is tentative due to the small sample size. We found that the risk of postpartum depression is affected both by mothers' temperament, and perinatal factors. Specifically, the two most strongly associated features with risk of postpartum depression for mothers with preterm infants are positive affect (r_s =-0.467, p < 0.001), and the Apgar score at 5 minutes after birth (r_s =-0.430, p<0.001). These two features have a negative association with postpartum depression, meaning that mothers with high positive affectivity (i.e. enthusiastic, energetic, confident) have a lower risk of developing postpartum depression. A lower Apgar score at 5 minutes after birth increases the risk of postpartum depression. Here, the Apgar score might be a proxy for subsequent events, such as neonatal morbidities or days spent in the NICU. Fear, another characteristic of adult temperament assessed by the Adult Temperament Questionnaire short form, has a strong positive association with risk of postpartum depression (r_s=0.399, p<0.01). On the other hand, extraversion (i.e. the tendency of primarily obtaining gratification from outside oneself) (r_s =-0.347, p < 0.01), attentional control (i.e. capacity to focus and shift attention when desired) ($r_s=-0.339$, p<0.01), and neutral sensitivity (i.e. detection of low intensity stimuli) (r_s =-0.307, p<0.05) have a strong negative association with postpartum depression. Moreover, risk of postpartum depression increases with decreasing gestational age (r_s =-0.318, p<0.01). The total days of intubation ($r_s=0.330$, p<0.01), as well as the total days of supplemental O_2 (r_s=0.323, p<0.01) that a preterm infant requires increase the risk of postpartum depression. These two exposures are proxies of respiratory co-morbidities of preterm birth,

which may cause distress to mothers. Finally, a past history of live births is positively associated with postpartum depression ($r_s=0.309$, p<0.01).

We found that the feature size giving the best results in terms of accuracy, sensitivity, specificity, and F1 score is 13 using the feature selection algorithm ReliefF expRank. The feature subset selected by the ReliefF expRank comprised past history of live births, bronchopulmonary dysplasia, extraversion, anxiety, total days of intubation and supplemental oxygen, Apgar score at 5 minutes after birth, effortful control, inhibitory control, positive affect, the infant's feeding at discharge, retinopathy of prematurity, and maternal asthma. We can conclude that extraversion, which is a characteristic of adult temperament, positive affect, as well as effortful control and inhibitory control are negatively correlated with the risk of postpartum depression. On the other hand, past history of anxiety and chronic diseases such as asthma may increase the risk of postpartum depression. Bronchopulmonary dysplasia, and retinopathy of prematurity are diseases that can result in significant morbidity for preterm neonates, causing distress to mothers. Furthermore, a lower Apgar score at 5 minutes after birth, and a higher number of days requiring intubation and supplemental oxygen increase the risk of postpartum depression; mothers who breastfeed their babies have a lower risk of depression compared to mothers whose babies drink infant formula. Finally, mothers with existing childcare responsibilities find it more challenging to cope after having a preterm infant compared with those who do not have pre-existing childcare duties. The performance of the RF resulted in out-of-sample balanced accuracy (median±IQR): 75.0 ± 16.7 , specificity: 100 ± 16.7 , and sensitivity: 66.7±16.7, and F1 score: 0.8±0.2.

Although we found some statistically strong correlations which gave some initial confidence that the statistical learning model would have a good chance of success, the performance of the RF was poor when taking into account the class imbalance in the investigated problem. One possible reason for the poor model performance is lack of statistical power: the sample size is not sufficiently large to effectively train the models. Another reason for poor model performance might be the fact that the given variables are insufficient to predict the response accurately. In addition, we need to bear in mind that a considerable number of features has been derived from questionnaires completed by the individuals themselves and therefore subjectivity in data collection may be another issue affecting the results.

We remark there are many contemporary advanced classifiers that can be used in order to map the features to the response, such as RF and Support Vector Machines (SVM). Here, in order to map the selected features to the response we only used RF and have not explored competing approaches (e.g. SVM) to optimize the classification performance further.

We envisage that a larger sample size may lead to a better and more generalizable model that may be more accurate in correctly assessing postpartum depression risk. Future work could potentially incorporate additional statistical machine learning algorithms which may improve prediction accuracy.

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