

Trajectory patterns and factors influencing perinatal fatigue among Chinese women from late pregnancy to six months after delivery

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Background. Perinatal fatigue among women is related to the clinical outcomes of mothers and infants. Perinatal fatigue changes over time, and the trajectory varies according to the predictors and outcomes of the mothers. This, however, has not been documented in any study.

Objectives. This study aimed to identify and characterize the trajectory patterns of perinatal fatigue among women from late pregnancy to six months after delivery.

Methods. We used growth mixture modeling to estimate the trajectory of perinatal fatigue at 28 gestational weeks (T0), 37 gestational weeks (T1), three days (T2), one week (T3), six weeks (T4), and six months (T5) after delivery with (n=1030). The Mann-Whitney U test and binary logistic regression were used to tie the selected trajectory classes to predictors and outcomes.

Results. There were two distinct patterns of perinatal fatigue in women: "persistently high" (11.1 %, n=114) and "persistently low" (88.9 %, n=916). Levels of perinatal fatigue among women in the "persistently high" group were higher than those in the "persistently low" group across the six measurements. Complications, fatigue at T0, and employment status in late pregnancy were all significant predictors of trajectories. Additionally, the "persistently high" group had a greater prevalence of difficult baby care and weight retention and a lower prevalence of exclusive breastfeeding.

Conclusions. Our study proved the heterogeneity and characteristics of perinatal fatigue among women. Future research should concentrate on developing intervention packages targeted at specific individuals in order to alleviate perinatal fatigue in women.

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Keywords. perinatal fatigue; trajectories; predictors; outcomes; growth mixture modeling

Introduction

Perinatal fatigue, which includes both pregnancy and postpartum fatigue, is one of the most frequently reported complaints among childbearing women^{1,2}. And perinatal fatigue is related to adverse clinical outcomes for women and infants. Prenatal fatigue could increase rates of pregnant morbidity³, cesarean section⁴, preterm birth, and even postpartum depression⁵. Postpartum fatigue may delay the initiation of breastfeeding⁶, reduce mother-infant attachments⁷ and even cause postpartum depression⁸. Collectively, comprehensive understandings of perinatal fatigue among women are needed.

However, longitudinal data on perinatal fatigue in women are quite scarce. Longitudinal studies have mainly focused on the levels and courses of perinatal fatigue among the population over time, which may obscure individual differences^{9,10,11}. Only one study, Kuo's investigation, has reported perinatal fatigue trajectories among 121 women from the late pregnancy to one week postpartum¹². Kuo's analysis clarified the heterogeneity of fatigue among women from pregnancy to postpartum periods. However, class 3 of the perinatal fatigue trajectories in Kuo's study only had nine women due to its limited sample sizes. And it only measured perinatal fatigue among women until one week postpartum, which shortened perinatal periods. Thus, it is necessary to gain understanding into whether and how perinatal fatigue trajectories emerge in larger samples of women throughout longer perinatal periods.

Investigating the factors that contribute to prenatal fatigue trajectories is also critical for intervention development. Few aspects of perinatal fatigue trajectories have been reported, though multiple factors are associated with the perinatal fatigue^{13,14}. In this study, the theoretical framework for factors influencing perinatal fatigue was according to Pugh's framework¹⁵. Pugh's framework used the Classification of Nursing Diagnosis definition of fatigue: "An overwhelming sustained sense of exhaustion and decreased capacity for physical and mental work." In this framework, related factors of perinatal fatigue were categorized into situational, physiologic, psychologic, and performance factors (**Figure 1**). Investigating the factors that influence

63 perinatal fatigue trajectories may provide to a better understanding of how to avoid or reduce
 64 perinatal fatigue in women.
 65 From late pregnancy to six months after birth, this prospective, longitudinal study assessed
 66 perinatal fatigue in women six times. The particular objectives of this study were to detect and
 67 define the trajectory patterns of perinatal fatigue in women from late pregnancy to six months
 68 after birth. Extending our understanding of the identified perinatal fatigue trajectory types and
 69 their related components may help to improve clinical management for perinatal fatigue.

Materials & Methods

Study Design and Settings

This study is a prospective, longitudinal study, conducted between December 2020 and September 2021. Participants were asked to fill out questionnaires six times, namely, 28 gestational weeks(T_0), 37 gestational weeks(T_1), three days postpartum(T_2), one week postpartum(T_3), six weeks postpartum(T_4), and six months postpartum(T_5). This work was approved by the Obstetrics and Gynecology Hospital attributed to Fudan University (No.2020-191).

Participants

We used the continuous enrollment method to find eligible participants between December 2020 and January 2021. Pregnant woman at (28+0, 29+6) weeks was scanned and recruited. The exclusion criteria were women with (a) diagnosed depression before; (b) severe health problems such as heart failure; (c) twins or multiples. All women signed informed consent.

Measurements

Perinatal fatigue was measured by the Postnatal Accumulated Fatigue Scale (PAFS). Professor Emi Mori firstly published the scale in 2016¹⁶. Each of the 13 items of this scale has a three-point rating from 0-3 (0 = rarely, 1 = sometimes and 3 = often). These 13 items are divided into three dimensions (physical, emotional, and cognitive), for example, "I feel unfocused" (cognitive). The total score ranges from 0 to 39, with higher scores indicating higher fatigue levels. This scale has good reliability and validity in the former study¹⁶. In this study, we obtained authorization from Professor Emi Mori regarding the use of the PAFS, and translated the Japanese version of PAFS into a Chinese version. The Chinese version of PAFS scale also has good reliability (Cronbach's $\alpha=0.823\sim0.908$) and validity ($KMO=0.801\sim0.91$) in this sample. Data of perinatal fatigue were collected six times from T_0 to T_5 .

Self-made questionnaires collected data of situational factors at T_0 and T_2 . At T_0 , we collected data of maternal age, parity(primipara, multipara), employment (employed, unemployed), an

education level (<college, college, >college), monthly family income (<¥10000, ¥10000~20000, >¥20000) and residency (urban, suburban). Also, we collected data of planned pregnancy (yes, no), prenatal training (yes, no), sleep duration per day (<six hours[hrs], six to eight hrs, >8hrs), and sleep quality at this time. Of them, sleep quality was evaluated by asking women, "Please score your sleep quality in the past 24 hours, ranging from 0 to 10, and 0 score means very dissatisfied, and 10 scores mean very satisfied." Besides, fetus gender (boys, girls) was obtained by medical records at T₂.

Self-made questionnaires were also used to collect data of physiologic factors at T₀ and T₁. Maternal height, prenatal weight, weight at T₀, gestation, complication, pain, and fatigue were collected at T₀. Of them, the pain was measured by asking one question: "Please score your pain in the past week, ranging from 0 to 10, and 0 score means no pain and 10 scores mean very painful." Based on these data, we calculated prenatal body mass index (BMI), BMI at T₀, weight gain at T₀ (weight at T₀ minus prenatal weight), and BMI gain at T₀ (BMI at T₀ minus prenatal BMI). And "T₁-T₀ fatigue" was equal to PAFS fatigue score at T₁ minus that at T₀, similar calculations were used for (T₂-T₁)/(T₃-T₂)/(T₄-T₃)/(T₅-T₄) Fatigue.

One maternal psychologic factor, anxiety, was measured at T₀ by asking participants: "How anxious are you in the past week? Choose a number between 0-10 to express the extent of your anxiety, 0 for no anxiety, 1 to 3 scores for mild anxiety, 4 to 6 scores for moderate anxiety, 7 to 9 scores for severe anxiety, and 10 scores for extreme anxiety."

Performance factors among women were regarded as outcomes of perinatal fatigue in this study, which were all collected at T₅, including difficult baby care (yes, no), exclusive breastfeeding (EBF; yes, no), and weight (kilogram, M±SD). Also, weight retention was defined as "yes" if weight at T₅ was more than prenatal weight, vice versa as "no."

Statistical analysis

Only data of participants who completed at least three follow-ups were included in the analysis.

Data analysis was carried out in the following two parts based on the two objectives of this study.

In the first part, we used growth mixture modeling (GMM) to identify the trajectory of perinatal fatigue. GMM is an advanced, person-centered analysis to identify heterogeneous subgroups that comprise distinct courses across time¹⁷. The time intervals were uneven from T_0 to T_5 , namely, 0 weeks (T_0-T_0), 8 weeks (T_1-T_0), 12 weeks (T_2-T_0), 13 weeks (T_3-T_0), 18 weeks (T_4-T_0), 36 weeks (T_5-T_0), respectively. Thus, the time score of T_0 to T_5 was fixed here at 0, 8, 12, 13, 18, 36, representing the equidistant time model. A robust maximum likelihood estimator was used for model estimation. Akaike Information Criterion (AIC), Bayesian Information Criterion (BIC), and Sample-Size Adjusted BIC (aBIC) were used as the model fit goodness statistics, with smaller values suggesting better model fit. Entropy was used to evaluate the accuracy of classification. The closer the entropy was to 1, the higher the class accuracy was. Likelihood Ratio Test (LRT) compared an n-class model with an n-1-class model, where a significant P value indicated a considerable improvement in the n-class model and thus rejected n-1-model rationally. In this study, the Lo-Mendell-Rubin (LMR) and Bootstrap Likelihood Ratio Test (BLRT) were also used for model comparison. Bayesian posterior probability was applied to export latent trajectory classes after a best-fit model was determined. In this part, GMM was performed with Mplus 8.3. In addition, a rank-sum test was used to examine differences in fatigue scores at six times and during five adjacent times between the trajectories. And $p < 0.05$ were preliminary remarked as positive.

In the second part, to characterize the trajectory of perinatal fatigue, the Mann-Whitney U test, and binary logistic regression were implemented. We calculated the Z-score of skewness and kurtosis for all continuous variables (Supplemental Table 1). Only gestation at baseline was normally distributed, but it does not have homogeneity of variance. Thus, the Mann-Whitney U test was used to compare differences between all variables, including continuous variables (such as maternal age) and categorical variables (such as monthly family income). Variables with $p < 0.05$ were preliminary remarked as positive between the trajectory classes. To further describe characteristics of perinatal fatigue trajectory, binary logistic regression (Forward: LR) was

148 applied. The latent trajectory classes as the dependent variable and positive potential predictors
 149 as independent variables in the regression model. And variables with $p < 0.05$ in the regression
 150 model were regarded as predictors of perinatal fatigue trajectory. In this part, performance
 151 factors were not included in the regression model because they were considered outcomes rather
 152 than potential predictors. SPSS version 23.0 was used for Mann-Whitney U and binary logistic
 153 regression.

Results

Of 1150 women who met initial eligibility criteria and consented to participation, a total of 1030 women who completed at least three follow-ups were included in this report. Cases included in each visit were 1009(T₀), 917(T₁), 970(T₂), 781(T₃), 563(T₄), and 779(T₅), respectively(**Figure 2**). The average age was 30.47 (SD 4.01) years in this study. Besides, 75.8% of these women were primiparas, 82.1% had a college degree, 80.9% were employed, and 87.1% had high monthly family income(\geq ¥ 10000). No significant difference of Basic information was found between included and non-included subjects (Supplemental Table 2).

Identifying Perinatal Fatigue Trajectories

Table 1 shows the GMM of one to five fatigue trajectory classes, including 1-class model, 2-class model, 3-class model, 4-class model, and 5-class model. All models can be supposed to obtain the best maximum likelihood. The best fit model is based on different model fit indices: (1)the AIC, BIC, and ABIC are smaller; (2)the entropy is closer to 1; (3)LMR and BLRT are both significant. Collectively, the 2-class model was the best fit model in this study. The mean intercept factor of Class 1(12.318) was higher than that of Class 2 (10.118), and the variance of the intercept factor between these two classes was 13.586 (p<0.001), indicating a relatively higher initial fatigue score in Class 1 than Class 2. The average slope factors of Class 1 and Class 2 were 0.306 (p<0.001) and -0.110 (p<0.001), indicating that Class 1's fatigue score increased significantly over time while Class 2's decreased significantly over time. In addition, the trajectories of fatigue scores were also visually shown in **Figure 3**. Thus, Class 1 and Class 2 were named as the "persistently high" group (11.1%; n=114) and "persistently low" group (88.90%; n=916), respectively.

Table 2 showed that fatigue scores had inconsistent growth tendency between different adjacent times. Firstly, an interesting finding was that the fatigue score in the "persistently high" group increased significantly faster than the "persistently low" group from T₀ to T₁ (p=0.001). And no significant differences in fatigue scores were found from T₁ to T₂, from T₂ to T₃, and from T₃ to

T₄. Combined **Table 1** with **Figure 1**, from T₀ to T₄, fatigue scores in both classes presented similar upward (from T₀ to T₃) and downward (from T₃ to T₄) trends. Oppositely, fatigue scores showed a strong upward trend from T₄ to T₅ in the "persistently high" group, whereas a solid downward trend in the "persistently low" group ($p < 0.001$). In addition, for the "persistently high" group, the peak fatigue score showed at T₅, followed by that at T₃. For the "persistently low" group, the peak of fatigue score showed at T₃.

Characteristics of Perinatal Fatigue Trajectories

Table 3 summarizes the two groups' descriptive statistics of related factors (situational, physiologic, psychologic, and performance factors). For situational factors, individuals in the "persistently high" group were less employed ($p = 0.002$) and had worse sleep quality ($p < 0.001$) at T₀ than those in the "persistently low" group. For physiologic factors, women in the "persistently high" group had smaller prenatal weight ($p = 0.014$), weight at T₀ ($p = 0.014$), BMI at T₀ ($p = 0.036$), more complication ($p < 0.001$), and severer fatigue at T₀ ($p < 0.001$) than those in "persistently low" group. For psychologic factors, more people in the "persistently high" group reported anxiety status with scores between 7-10 at T₀ than the "persistently low" group did ($p < 0.001$). Binary logistic regression was used to analyze the relationships between potential factors and the perinatal fatigue trajectory. The regression model was summarized in **Table 4**. In the model, perinatal fatigue trajectory classes were the dependent variable and coded as 1 = "persistently high" group and 0 = "persistently low" group. And all factors significantly differed between the two classes were regarded as independent variables. But a high correlation was found between fatigue at T₀ and sleep quality ($r = 0.38$, $p < 0.001$), fatigue at T₀ and anxiety ($r = 0.36$, $p < 0.001$), BMI at T₀ and prenatal weight ($r = 0.79$, $p < 0.001$), and BMI at T₀ and Weight at T₀ ($r = 0.87$, $p < 0.001$). Thus, only employment (1 = "employed" and 0 = "unemployed"), complication (1 = "yes" and 0 = "no"), fatigue at T₀, and BMI at T₀ from 878 cases were introduced into the final model. The χ^2 test of the final model was significant ($P < 0.001$), and the Hosmer and Lemeshow test indicated an acceptable model fit with $p = 0.417$. As a result, the final model showed three

206 predictors of perinatal fatigue trajectories, i.e. complication (odds ratio [OR], 2.779, 95%
 207 confidence interval [CI], 1.641~4.706), fatigue at T0 (OR, 1.086, 95% CI, 1.042~1.132) and
 208 employment (OR, 0.544; 95% CI, 0.310~0.954).
 209 In addition, **Table 3** also showed differences in the performance factors between the two groups.
 210 The percentages of patients with difficult baby care (36.8% VS 16.9%; $p < 0.001$) and weight
 211 retention (56.8% VS 40.9%; $p = 0.001$) were significantly higher in the "persistently high" group
 212 than the other group at T₅. Also, fewer people reported exclusive breastfeeding (EBF, 32.6% VS
 213 47%; $p = 0.002$) in the "persistently high" group than the other group at T₅.

DISCUSSIONS

Our study addressed previously unresolved gaps in the literature by identifying and describing the perinatal fatigue trajectory in a large sample from late pregnancy to six months postpartum. The findings of this study are critical for developing therapies to alleviate perinatal fatigue and improve mother and child clinical outcomes. We discovered two unique perinatal fatigue trajectories, namely "persistently high" (class 1, 11.1%) and "persistently low"(class 2, 88.9%). Additionally, we were the first to identify predicted markers for the perinatal fatigue trajectory, such as complication, fatigue at T₀, and employment. Additionally, women in the "persistently high" group experienced more adverse clinical outcomes than those in the "persistently low" group, including greater rates of difficult baby care and weight retention and lower rates of exclusive breastfeeding at T₅.

The identification of two distinct trajectories of prenatal fatigue in women confirmed the variability of perinatal fatigue. Though few research have shown this in pregnant women, prior investigations have identified two distinct fatigue trajectories in other populations, such as patients with cancer ¹⁸ and systemic sclerosis¹⁹. To our knowledge, only Kuo's study has investigated this in perinatal women, identifying three distinct trajectories (class 1, 44.6 %; class 2, 48.0 %; and class 3, 3.7%) of perinatal fatigue from late pregnancy to one week postpartum in 121 women¹². Kuo's study detected one additional trajectory of perinatal fatigue in women than ours, which could be explained by the difference in measurement frequency, follow-up time, and sample size. Further research could be undertaken to confirm the trajectories of perinatal fatigue in women. Additionally, both our study and Kuo's study discovered that around one-tenth (11.1 % versus 7.4 %) of women were at high risk of experiencing extreme fatigue during their perinatal periods. Clinical personnel should identify and monitor these high-risk women more closely.

Notably, levels of perinatal fatigue were significantly higher in women classified as "persistently high" than in women classified as "persistently low" at all six time points from late pregnancy to

six months after delivery, which was consistent with prior study in other patients^{19,20,21}. Additionally, the general level of prenatal fatigue grew dramatically with time in the "persistently high" group, whereas it decreased significantly in the "persistently low" group. This characteristic revealed that fatigue could be closely related to specific internal components. Two time points have been identified for decreasing perinatal fatigue in women. One was one week postpartum for women classified as "persistently low." This conclusion is consistent with previous research emphasizing the critical role of one week postpartum in preventing perinatal fatigue^{22,23}. The other was six months after birth for women classified as having "persistently high" blood pressure. On the other hand, Taylor's study found that perinatal fatigue reached a minimum six months following delivery²⁴. And, in this study, the lowest level of perinatal fatigue was also six months after delivery in the "persistently low" group. This discrepancy may be explained by Taylor's study's analysis of the peak of perinatal fatigue using trends rather than trajectories. Thus, for the majority of women, the peak of perinatal weariness occurred one week after delivery, whereas for others, it occurred six months later. Future research could examine the factors that contribute to fatigue in women one week and six months following delivery. In our investigation, a complication was identified as a risk factor for developing an elevated perinatal fatigue pattern. Women who experienced complications during late pregnancy, such as GDM²⁵ or hypertension²⁶, reported higher level of perinatal fatigue. Complications have also been implicated as risk factors for fatigue in pregnant women²⁷ and postpartum women²⁸. Pathways for preventing or alleviating problems during late pregnancy should be researched further. Another intriguing discovery was that exhaustion during late pregnancy may have an effect on how subsequent perinatal fatigue develops over time. Women who had increased fatigue throughout late pregnancy were more likely to fall into the "persistently high" category. Although this discovery has not been replicated in other studies, a similar pattern has been observed in prenatal depression²⁹. Thus, health care providers could identify women at risk of

extreme exhaustion throughout the perinatal period as early as late pregnancy. Additionally, future research could develop predictive models to validate the prediction of perinatal fatigue in women based on exhaustion during late pregnancy.

Employment status was also associated with perinatal fatigue trajectories. This result suggested that women whose employment ratio was "persistently high" had a lower employment ratio during late pregnancy. Similarly to our employment finding, Chien's study found that unemployed women experienced considerably higher levels of weariness than other women during pregnancy⁴. This difference could be explained by job insecurity, which has been shown to impair psychological resilience³⁰. As a result, women should be encouraged to seek employment rather than evade it in late pregnancy.

Additionally, to examine determinants of perinatal fatigue trajectories in women, we evaluated variations in performance parameters such as challenging baby care ratios, EBF, and weight retention at six months postpartum across the two trajectories. We discovered that the "persistently high" group had a larger proportion of challenging baby care than the other group, which is consistent with previous reports^{7, 31}. Both of these investigations, however, were cross-sectional. Thus, our study was the first to reveal the detrimental effect of perinatal fatigue on six-month-old newborn care.

Does fatigue impact breastfeeding in women during their reproductive years? This issue has dogged health care providers for 15 years^{6,32}. At six months postpartum, our study discovered that the ratio of exclusive breastfeeding was considerably lower in the "persistently high" group than in the "persistently low" group. Despite the cross-sectional nature of the study, earlier research has also demonstrated a detrimental relationship between postpartum fatigue and breastfeeding outcomes^{33,34}. Logically, a higher level of postpartum fatigue among women was related to delayed breastfeeding initiation³⁵, decreased self-efficacy²³, and increased breastfeeding difficulties³⁶, all of which finally result in the termination of exclusive breastfeeding³⁷. Our study was the first to establish a link between the prenatal fatigue trajectory

and the outcomes of exclusive breastfeeding six months after delivery, using a different study design and data analysis method than previous studies.

In addition, women need to escape weight retention^{38,39,40}. At six months postpartum, 56.8 percent of women in the "persistently high" group experienced weight retention, compared to 40.9 percent of women in the "persistently low" group. To our knowledge, no other study has examined the association between perinatal fatigue and weight retention in postpartum women. Additional research is required to determine the association between fatigue and weight retention. Several limitations should be highlighted regarding this investigation. To begin, our study stopped collecting data on perinatal fatigue six months after birth. Fatigue may occur in women who deliver more than a year after delivery^{41,42}. A longer duration is necessary for examining the trajectories of perinatal fatigue in women. Second, the exact instrument used to assess perinatal fatigue was developed in 2016 and is not yet widely available. There is a need for reports on the implementation of the PAFS scale in perinatal mothers from diverse cultural backgrounds. Third, anxiety was the sole psychological element during the perinatal period, according to Pugh's theoretical model of perinatal fatigue. As a result, we focused only on anxiety as a psychological component of perinatal fatigue. And it was quantified using a single question rather than a predefined scale. Other psychologic variables, such as depression, should also be considered in future studies. Fourth, due to the researchers' limited time, only 574 women completed the six-week postpartum follow-up. Additionally, 92.5% of our participants were college or university graduates. Our findings should be reproduced in lower-education populations.

This study has a few implications for perinatal care research and practice. For research purposes, future empirical investigations could validate and change the perinatal fatigue trajectory classes among women from many cultural backgrounds. Additionally, more predictive models should be developed to investigate and illustrate prenatal fatigue predictors. This will allow for future investigation of the characteristics of prenatal fatigue and the development of more effective therapies targeting sub-populations for lowering individual perinatal fatigue. Additionally, future

research should examine the causal links between prenatal fatigue and clinical outcomes such as rates of difficult baby care, exclusive breastfeeding, and weight retention, thereby filling in the gaps in the data. In clinical practice, health care practitioners could undertake timely examinations of women early in their late pregnancy and screen out high-risk women who are at danger of experiencing extreme fatigue throughout the postnatal period.

To summarize, this study demonstrates the heterogeneity of perinatal fatigue in women by distinguishing two distinct perinatal fatigue trajectories: "persistently high" and "persistently low." Additionally, this study examined the characteristics of the two perinatal fatigue trajectories by finding three late-pregnancy predictors: complication, fatigue, and employment. Future research should focus on developing customized intervention packages based on these findings in order to alleviate women's perinatal fatigue.

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References

1. Kilic M, Eryilmaz G. The determination of women's fatigue levels in the third trimester of pregnancy and in the postpartum period. *Healthmed*. 2011;5(4):826-830.
2. Liu N, Wang J, Chen D dan, Sun W jia, Li P, Zhang W. Effects of exercise on pregnancy and postpartum fatigue: A systematic review and meta-analysis. *Eur J Obstet Gynecol Reprod Biol*. 2020;253:285-295. doi:10.1016/j.ejogrb.2020.08.013
3. Luke B, Avni M, Min L, Misiunas R. Work and pregnancy: the role of fatigue and the "second shift" on antenatal morbidity. *Am J Obstet Gynecol*. 1999;181(5 Pt 1):1172-1179. doi:10.1016/s0002-9378(99)70103-1
4. Chien LY, Ko YL. Fatigue during pregnancy predicts caesarean deliveries. *J Adv Nurs*. 2004;45(5):487-494. doi:10.1046/j.1365-2648.2003.02931.x
5. Newman RB, Goldenberg RL, Moawad AH, et al. Occupational fatigue and preterm premature rupture of membranes. National Institute of Child Health and Human Development Maternal-Fetal Medicine, Units Network. *Am J Obstet Gynecol*. 2001;184(3):438-446. doi:10.1067/mob.2001.110312
6. Saidi L, Godbout P. Study of two indicators of difficulty in initiating maternal breastfeeding : maternal fatigue and suboptimal infant breastfeeding behaviour. *Rech Soins Infirm*. 2016;(125):32-45. doi:10.3917/rsi.125.0032
7. Lai YL, Hung CH, Stocker J, Chan TF, Liu Y. Postpartum fatigue, baby-care activities, and maternal-infant attachment of vaginal and cesarean births following rooming-in. *Appl Nurs Res ANR*. 2015;28(2):116-120. doi:10.1016/j.apnr.2014.08.002
8. Wilson N, Lee JJ, Bei B. Postpartum fatigue and depression: A systematic review and meta-analysis. *J Affect Disord*. 2019;246:224-233. doi:10.1016/j.jad.2018.12.032
9. Cheng CY, Pickler RH. Perinatal stress, fatigue, depressive symptoms, and immune modulation in late pregnancy and one month postpartum. *ScientificWorldJournal*. 2014;2014:652630. doi:10.1155/2014/652630
10. Cheng CY, Chou YH, Wang P, Tsai JM, Liou SR. Survey of trend and factors in perinatal maternal fatigue. *Nurs Health Sci*. 2015;17(1):64-70. doi:10.1111/nhs.12149
11. Iwata H, Mori E, Sakajo A, Aoki K, Maehara K, Tamakoshi K. Course of maternal fatigue and its associated factors during the first 6 months postpartum: a prospective cohort study. *Nurs Open*. 2018;5(2):186-196. doi:https://doi.org/10.1002/nop2.130

12. Kuo SY, Yang YL, Kuo PC, Tseng CM, Tzeng YL. Trajectories of Depressive Symptoms and Fatigue Among Postpartum Women. *Jognn-J Obstet Gynecol Neonatal Nurs*. 2012;41(2):216-226. doi:10.1111/j.1552-6909.2011.01331.x
13. Badr HA, Zauszniewski JA. Meta-analysis of the predictive factors of postpartum fatigue. *Appl Nurs Res*. 2017;36:122-127. doi:10.1016/j.apnr.2017.06.010
14. Yehia DBM, Malak MZ, Al-Thwabih NN, et al. Psychosocial factors correlate with fatigue among pregnant women in Jordan. *Perspect Psychiatr Care*. 2020;56(1):46-53. doi:10.1111/ppc.12372
15. Pugh L, Milligan R. A Framework for the Study of Childbearing Fatigue. *Adv Nurs Sci*. 1993;15(4):60-70. doi:10.1097/00012272-199306000-00007
16. Tsuchiya M, Mori E, Sakajo A, Iwata H, Maehara K, Tamakoshi K. Cross-sectional and longitudinal validation of a 13-item fatigue scale among Japanese postpartum mothers. *Int J Nurs Pract*. 2016;22:5-13. doi:10.1111/ijn.12434
17. Feldman BJ, Masyn KE, Conger RD. New Approaches to Studying Problem Behaviors: A Comparison of Methods for Modeling Longitudinal, Categorical Adolescent Drinking Data. *Dev Psychol*. 2009;45(3):652-676. doi:10.1037/a0014851
18. Bødtcher H, Bidstrup PE, Andersen I, et al. Fatigue trajectories during the first 8 months after breast cancer diagnosis. *Qual Life Res*. 2015;24(11):2671-2679. doi:10.1007/s11136-015-1000-0
19. Willems LM, Kwakkenbos L, Vonk MC, van den Hoogen FHJ, Vlieland TPMV, van den Ende CHM. Three-year trajectories of disability and fatigue in systemic sclerosis: a cohort study. *Clin Exp Rheumatol*. 2017;35(4):S48-S55.
20. Li H, Marsland AL, Conley YP, Sereika SM, Bender CM. Genes Involved in the HPA Axis and the Symptom Cluster of Fatigue, Depressive Symptoms, and Anxiety in Women With Breast Cancer During 18 Months of Adjuvant Therapy. *Biol Res Nurs*. 2020;22(2):277-286. doi:10.1177/1099800419899727
21. Bean HR, Diggins J, Ftanou M, Weihs KL, Stanton AL, Wiley JF. Insomnia and Fatigue Symptom Trajectories in Breast Cancer: A Longitudinal Cohort Study. *Behav Sleep Med*. Published online January 20, 2021:1-14. doi:10.1080/15402002.2020.1869005
22. Henderson J, Alderdice F, Redshaw M. Factors associated with maternal postpartum fatigue: an observational study. *Bmj Open*. 2019;9(7):e025927. doi:10.1136/bmjopen-2018-025927

- 394 23. Fata S, Atan SU. The Relationship between Fatigue and Breastfeeding Self-efficacy. Niger J
395 Clin Pract. 2018;21(11):1408-1414. doi:10.4103/njcp.njcp_366_17
- 396 24. Taylor J, Johnson M. How women manage fatigue after childbirth. Midwifery.
397 2010;26(3):367-375. doi:10.1016/j.midw.2008.07.004
- 398 25. Zhang J, Chi H, Xiao H, et al. Interleukin 6 (IL-6) and Tumor Necrosis Factor α (TNF- α)
399 Single Nucleotide Polymorphisms (SNPs), Inflammation and Metabolism in Gestational
400 Diabetes Mellitus in Inner Mongolia. Med Sci Monit Int Med J Exp Clin Res. 2017;23:4149-
401 4157. doi:10.12659/msm.903565
- 402 26. Li Y, Wang Y, Ding X, Duan B, Li L, Wang X. Serum Levels of TNF- α and IL-6 Are
403 Associated With Pregnancy-Induced Hypertension. Reprod Sci Thousand Oaks Calif.
404 2016;23(10):1402-1408. doi:10.1177/1933719116641760
- 405 27. Mortazavi F, Borzoe F. Fatigue in Pregnancy. Sultan Qaboos Univ Med J. 2019;19(1):e44-
406 e50. doi:10.18295/squmj.2019.19.01.009
- 407 28. Taylor J, Johnson M. The role of anxiety and other factors in predicting postnatal fatigue:
408 From birth to 6 months. Midwifery. 2013;29(5):526-534. doi:10.1016/j.midw.2012.04.011
- 409 29. Jacques N, Mesenburg MA, Matijasevich A, et al. Trajectories of maternal depressive
410 symptoms from the antenatal period to 24-months postnatal follow-up: findings from the 2015
411 Pelotas birth cohort. BMC Psychiatry. 2020;20(1):233. doi:10.1186/s12888-020-02533-z
- 412 30. Heinz AJ, Meffert BN, Halvorson MA, Blonigen D, Timko C, Cronkite R. Employment
413 characteristics, work environment, and the course of depression over 23 years: Does employment
414 help foster resilience? Depress Anxiety. 2018;35(9):861-867. doi:10.1002/da.22782
- 415 31. Henderson J, Alderdice F, Redshaw M. Factors associated with maternal postpartum fatigue:
416 an observational study. Bmj Open. 2019;9(7):e025927. doi:10.1136/bmjopen-2018-025927
- 417 32. Callahan S, Sejourne N, Denis A. Fatigue and breastfeeding: An inevitable partnership? J
418 Hum Lact. 2006;22(2):184-187. doi:10.1177/0890334406286972
- 419 33. Brown CRL, Dodds L, Legge A, Bryanton J, Semenik S. Factors influencing the reasons why
420 mothers stop breastfeeding. Can J Public Health-Rev Can Sante Publique. 2014;105(3):E179-
421 E185. doi:10.17269/cjph.105.4244
- 422 34. Aldalili AYA, El Mahalli AA. Research Title: Factors Associated with Cessation of
423 Exclusive Breastfeeding. J Multidiscip Healthc. 2021;14:239-246. doi:10.2147/JMDH.S277819

35. Heinig MJ. Addressing Maternal Fatigue: A Challenge to In-hospital Breastfeeding Promotion. *J Hum Lact*. 2010;26(3):231-232. doi:10.1177/0890334410378975
36. Cooklin AR, Amir LH, Nguyen CD, et al. Physical health, breastfeeding problems and maternal mood in the early postpartum: a prospective cohort study. *Arch Womens Ment Health*. 2018;21(3):365-374. doi:10.1007/s00737-017-0805-y
37. Gianni ML, Bettinelli ME, Manfra P, et al. Breastfeeding Difficulties and Risk for Early Breastfeeding Cessation. *Nutrients*. 2019;11(10). doi:10.3390/nu11102266
38. Durst J, Tuuli M, Stout M, Macones G, Cahill A. Postpartum weight retention: impact on perinatal outcomes in subsequent pregnancy. *Am J Obstet Gynecol*. 2015;212(1):S176-S177. doi:10.1016/j.ajog.2014.10.375
39. Martin J, MacDonald-Wicks L, Hure A, Smith R, Collins CE. Reducing Postpartum Weight Retention and Improving Breastfeeding Outcomes in Overweight Women: A Pilot Randomised Controlled Trial. *Nutrients*. 2015;7(3):1464-1479. doi:10.3390/nu7031464
40. Sharkey KM, Boni GM, Quattrucci JA, Blatch S, Carr SN. Women with postpartum weight retention have delayed wake times and decreased sleep efficiency during the perinatal period: a brief report. *Sleep Health*. 2016;2(3):225-228. doi:10.1016/j.sleh.2016.05.002
41. Parks PL, Lenz ER, Milligan RA, Han HR. What happens when fatigue lingers for 18 months after delivery? *J Obstet Gynecol Neonatal Nurs JOGNN*. 1999;28(1):87-93. doi:10.1111/j.1552-6909.1999.tb01969.x
42. Wilson N, Wynter K, Fisher J, Bei B. Related but different: distinguishing postpartum depression and fatigue among women seeking help for unsettled infant behaviours. *Bmc Psychiatry*. 2018;18:309. doi:10.1186/s12888-018-1892-7
43. Dokuhaki S, Dokuhaki F, Akbarzadeh M. The relationship of maternal anxiety, positive and negative affect schedule, and fatigue with neonatal psychological health upon childbirth. *Contracept Reprod Med*. 2021;6(1):12. doi:10.1186/s40834-021-00155-8

Figure 1

Framework for the study of mother's fatigue during the childbearing experience.

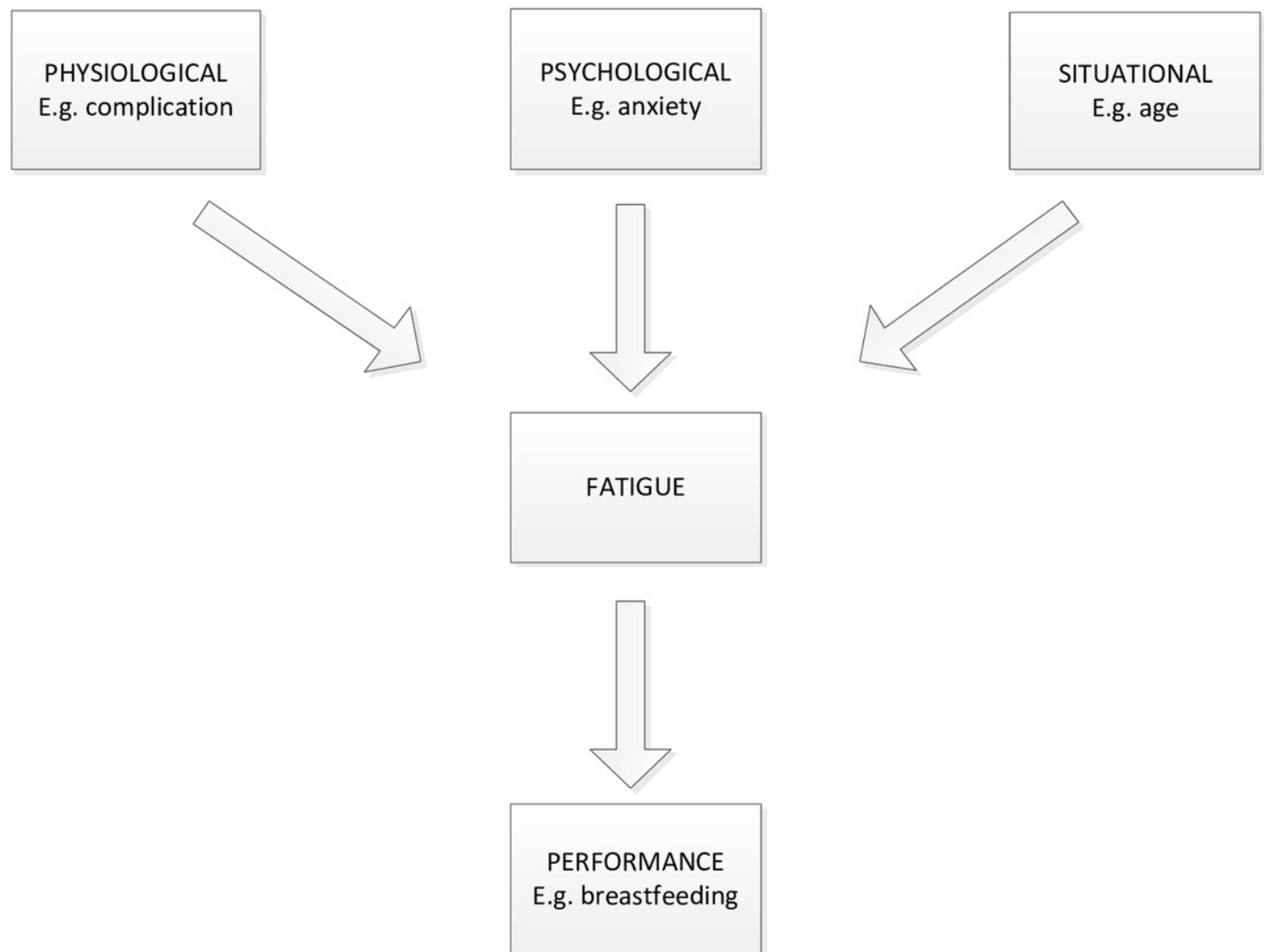


Figure 2

A flowchart of the participants in this study.

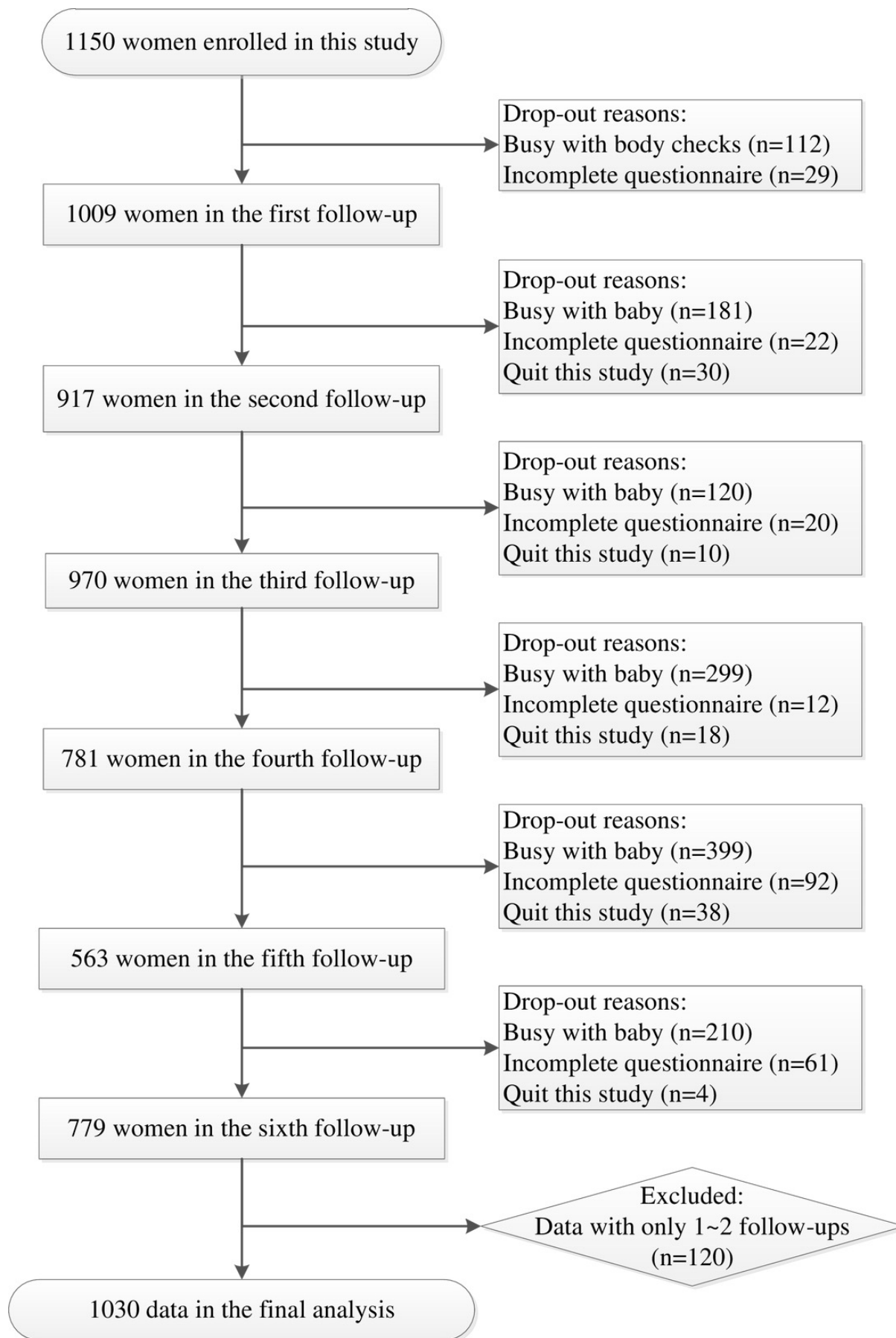


Figure 3

Two-class model of perinatal fatigue trajectory among women.

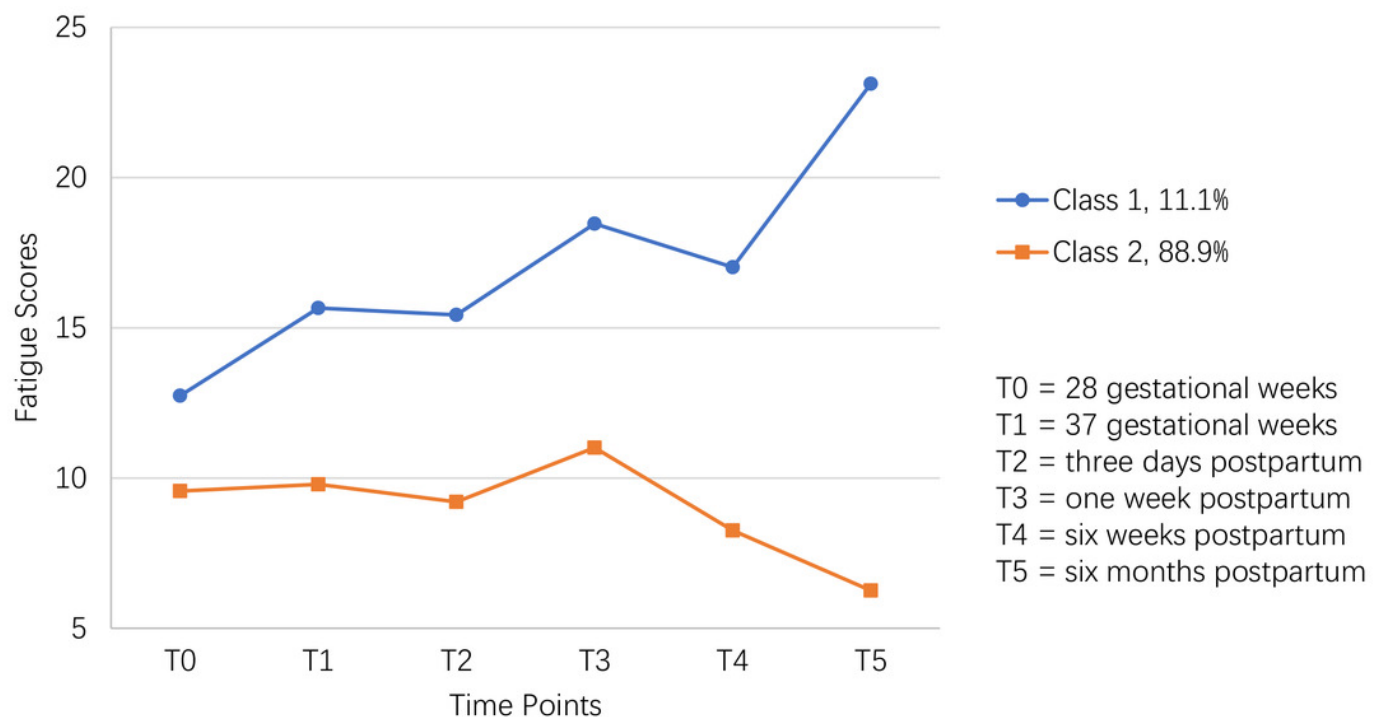


Table 1(on next page)

Fit indices for one to five fatigue trajectory classes of GMM.

1 **Table 1 Fit indices for one to five fatigue trajectory classes of GMM.**

C	k	AIC	BIC	aBIC	E	P of LMR	P of BLRT	class probability
1	11	33257.23	33311.54	33276.61	—	—	—	1
2	14	33022.68	33091.80	33047.33	.865	.000	.0000	.908/.092
3	17	32989.33	33073.26	33019.27	.821	.459	.0000	.850/.096/.053
4	20	32964.37	33063.11	32999.59	.831	.050	.0000	.830/.101/.057/.012
5	23	32951.67	33065.23	32992.18	.798	.360	.0000	.772/.109/.097/.012/.011

2 *Footnotes.* C = number of classes; k = number of free parameters; LL = loglikelihood; AIC = Akaike
 3 Information Criterion; BIC=Bayesian Information Criterion, aBIC = sample size adjusted BIC; E= Entropy;
 4 LMR = Lo-Mendell-Rubin test; BLRT = Bootstrap Likelihood Ratio Test; class probability=proportion of
 5 sample classification.

6

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Table 2(on next page)

Fatigue scores at six times and during five adjacent times.

Table 2 Fatigue scores at six times and during five adjacent times.

Variables	Persistently High (m, SD, M)	Persistently Low (m, SD, M)	U	Z	p
Fatigue at T ₀	12.74(6.312), 12.00	9.57(5.455), 9.00	55314.500	4.937	<0.001
Fatigue at T ₁	15.66(9.53), 15.50	9.79(7.373), 8.00	46577.500	5.399	<0.001
Fatigue at T ₂	15.43(7.894), 14.00	9.2(6.813), 7.00	59067.000	7.509	<0.001
Fatigue at T ₃	18.47(9.336), 18.00	11.01(7.483), 10.00	38128.500	6.703	<0.001
Fatigue at T ₄	17.02(10.846), 18.00	8.26(6.323), 6.00	23391.500	6.071	<0.001
Fatigue at T ₅	23.12(4.518), 22.00	6.26(3.993), 6.00	52148.000	14.188	<0.001
T ₁ -T ₀ Fatigue	3.28(8.864), 2.00	0.38(7.602), 0.00	38087.500	2.651	0.008
T ₂ -T ₁ Fatigue	0.05(11.23), -1.00	-0.73(7.713), 0.00	33258.000	0.131	0.896
T ₃ -T ₂ Fatigue	2.86(11.28), 3.00	1.62(8.839), 1.00	23955.000	0.778	0.436
T ₄ -T ₃ Fatigue	-0.83(14.432), 0.00	-3.06(8.791), -4.00	16427.500	0.456	0.648
T ₅ -T ₄ Fatigue	9.62(10.012), 11.00	-2.1(6.504), -1.00	12552.500	6.947	<0.001

Footnotes. Mann-Whitney U test. m=mean, SD=standard deviation, M=median, T₀=28 gestational weeks, T₁=37 gestational weeks, T₂=three days postpartum, T₃=one week postpartum, T₄=six weeks postpartum, T₅=six months postpartum. T₁-T₀ Fatigue=fatigue at T₁ minus that at T₀. (T₂-T₁)/(T₃-T₂)/(T₄-T₃)/(T₅-T₄) Fatigue were named in similar way.

Table 3(on next page)

Descriptive statistics of factors between the trajectory classes.

1 **Table 3 Descriptive statistics of factors between the trajectory classes.**

Variables	Persistently High	Persistently Low	U	Z	p
Number of participant, n(%)	95(9.2)	935(90.8)	-	-	-
Situational Factors					
Age, m(SD), M, y	30.21(3.57), 30	30.45(3.66), 30	40745.5	-0.959	0.337
Parity, n(%)			42199.0	-1.013	0.311
Primipara	76(80.0)	702(75.1)			
Multipara	19(20.0)	230(24.6)			
Missing	0(0.0)	3(0.3)			
Employment, n(%)			46473.0	3.049	0.002
Employed	62(65.3)	749(80.1)			
Unemployed	28(29.5)	163(17.4)			
Missing	5(5.2)	23(2.5)			
Education Level, n(%)			44252.0	0.506	0.613
<college	7(7.4)	73(7.8)			
college	16(16.8)	175(18.7)			
>college	71(74.7)	671(71.8)			
Missing	1(1.1)	16(1.7)			
Monthly family Income, n(%)			40010.5	-0.489	0.625
<¥ 10,000	12(12.6)	118(12.6)			
¥ 10,000~20,000	47(49.5)	455(48.7)			
>¥ 20,000	31(32.6)	342(36.6)			
Missing	5(5.3)	20(2.1)			
Residency, n(%)			41431.5	-1.824	0.068
Urban	93(97.9)	891(95.3)			
Suburban	0(0)	32(3.4)			
Missing	2(2.1)	12(1.3)			
Planned Pregnancy, n(%)			39788.5	-1.543	0.123
Yes	59(62.1)	646(69.1)			
No	35(36.8)	271(29.0)			
Missing	1(1.1)	18(1.9)			
Prenatal Training, n(%)			39422.0	-1.451	0.147
Yes	17(17.9)	233(24.9)			
No	75(78.9)	687(73.5)			
Missing	3(3.2)	15(1.6)			
Sleep Duration, n(%), hours			44526.0	1.898	0.058
<six	9(9.5)	97(10.4)			
Six to eight	53(55.8)	651(69.6)			
>eight	26(27.4)	173(18.5)			
Missing	7(7.4)	14(1.5)			
Sleep quality, m(SD),M	7.00(1.10), 7	7.48(1.19), 8	31530.5	-4.252	<0.001
Fetus Gender, n(%)			37380.5	1.222	0.222
Boys	39(41.1)	425(45.5)			
Girls	47(49.5)	388(41.5)			
Missing	9(9.5)	122(13.0)			
Physiologic Factors					
Height, m(SD), M, cm	161.99(4.79),1.62	162.76(4.93),1.63	39954.5	-1.332	0.183
Prenatal Weight, m(SD), M	53.71(7.97),52.00	55.78(8.24),55.00	35223.0	-2.456	0.014
Prenatal BMI, m(SD), M	20.44(2.64),20.19	21.04(2.83),20.70	36474.5	-1.967	0.050
Weight at T ₀ , m(SD), M	62.89(8.22),61.00	65.20(8.66),65.00	35899.0	-2.487	0.013

BMI at T ₀ , m(SD), M	23.93(2.66), 23.63	24.59(2.96), 24.43	36905.0	-2.094	0.036
Weight Gain at T ₀ , m(SD), M	9.05(2.83), 9.00	9.43(3.48), 9.05	38264.5	-1.132	0.258
BMI Gain at T ₀ , m(SD), M	3.45(1.06), 3.61	3.56(1.31), 3.54	38957.0	-0.849	0.396
Gestation, m(SD), M	29.07(0.70), 29.14	29.18(0.74), 29.14	40055.5	-0.882	0.378
Complication, n(%)			28058.5	-3.877	<0.001
YES	27(28.4)	139(14.9)			
NO	53(55.8)	710(75.9)			
Missing	15(15.8)	86(9.2)			
Pain, m(SD), M	2.54(2.70), 1.50	2.03(2.28), 2.00	36871.0	1.264	0.206
Fatigue at T ₀ , m(SD), M	12.74(6.312), 12.00	9.57(5.455), 9.00	55314.5	4.937	<0.001
Psychologic Factor					
Anxiety, n(%)			46235.0	5.419	<0.001
One to three	12(12.6)	243(26.0)			
Four to six	12(12.6)	230(24.6)			
Seven to nine	17(17.9)	194(20.7)			
Ten	41(43.2)	168(18.0)			
Missing	13(13.7)	100(10.7)			
Maternal Outcomes					
Difficult Baby Care, n(%)			32522.0	4.706	<0.001
YES	35(36.8)	158(16.9)			
NO	39(41.1)	546(58.4)			
Missing	21(22.1)	231(24.7)			
Exclusive Breastfeeding, n(%)			20217.5	-3.102	0.002
YES	31(32.6)	439(47.0)			
NO	40(42.1)	263(28.1)			
Missing	24(25.3)	233(24.9)			
Weight at T ₅ , m(SD), M	57.48(7.95), 56.50	56.98(8.34), 55.00	26815.5	0.844	0.399
Weight Retention at T ₅ , n(%)			18802.5	-3.281	0.001
YES	55(57.9)	380(42.0)			
NO	15(15.8)	293(30.0)			
Missing	25(26.3)	262(28.0)			

Footnotes. Mann-Whitney U test. m=mean, SD=standard deviation, M=median, n=number, %=percentage, y=year(s), cm=centimeters(s), h=hour(s), BMI=body mass index, T₀=28 gestational weeks, T₁=37 gestational weeks, T₅=six months postpartum.

Table 4(on next page)

Binary Logistic Regression for Class Membership.

1 **Table 4 Binary Logistic Regression for Class Membership.**

Variable	B	SE	W	df	P	OR	95%CI OR
Complication	1.022	0.269	14.453	1	<0.001	2.779	1.641~4.706
Fatigue at T ₀	0.083	0.021	15.553	1	<0.001	1.086	1.042~1.132
Employment	-0.609	0.287	4.508	1	0.034	0.544	0.310~0.954
constant	-3.101	0.356	75.992	1	<0.001	0.045	-

2 *Footnotes.* Binary Logistic Regression. B=regression coefficient, SE=standard error; W=ward,
 3 df=degree of freedom; OR=odd ratio; CI=confidence interval.