

Poor preoperative sleep quality is a risk factor for severe postoperative pain after breast cancer surgery

A prospective cohort study

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Abstract

The aim of this study was to assess the effect of preoperative sleep quality on acute postoperative pain in breast cancer patients.

The Pittsburgh Sleep Quality Index questionnaire (PSQI) was used to assess the overall sleep status of women scheduled for unilateral modified radical mastectomy in the past month. Based on the responses, patients were allocated to good sleep group or poor sleep group. Postoperatively, acute pain was assessed using the numerical rating score in the first 24 hours; in addition, the requirement of analgesics and the incidence of postoperative complications were recorded.

A total of 108 breast surgery patients were enrolled. Based on the PSQI results, 55 (51%) patients were allocated to poor sleep group and 53 (49%) to good sleep group. Pain scores were similar in the 2 groups at the end of surgery ($P = .589$); however, poor sleep group reported higher postoperative pain scores than the good sleep group at 2 ($P = .002$), 6 ($P < .001$), 12 ($P < .001$), and 24 ($P = .002$) hours after surgery. The incidence of severe pain in the poor sleep group was higher than that in the good sleep group (27% vs 8%, $P = .018$), and the ratio of participants who required rescued analgesics was greater in the poor sleep group (52% vs 22%, $P = .002$). In addition, patients with poor sleep quality had more postoperative complications and longer hospital stay.

In this study, breast cancer patients with poor preoperative sleep quality reported more severe postoperative pain, required more analgesics, experienced more complications, and had longer hospital stay.

Abbreviations: NRS = numerical rating score, PACU = postanesthesia care unit, PONV = postoperative nausea and vomiting, PSQI = Pittsburgh Sleep Quality Index, SAS = Self-rating Anxiety Scale, SDS = Self-rating Depression Scale.

Keywords: breast cancer surgery, numerical rating score, postoperative pain, sleep quality

1. Introduction

Acute postoperative pain after surgery is associated with increased postoperative complications, lower patient satisfaction, prolonged hospitalization, and greater medical expenditure^[1,2]; in addition, it also has an effect on the incidence of chronic pain and postoperative consumption of analgesics at home, especially

in patients receiving breast surgery, coronary artery bypass surgery, and thoracotomy.^[3,4] Better management of postoperative pain complies with the notion of comfort medicine and enhances postoperative recovery; in addition, it also decreases the incidence of persistent postoperative pain and helps improve the long-term quality of life.^[5] Currently, several modalities are available for effective management of postoperative pain^[6,7]; however, identification of patients who are more sensitive to pain and those who are likely to experience severe postoperative pain is still difficult.^[8] Therefore, it is important to identify the determinants of acute postoperative pain in order to improve the protocol for postoperative pain management.

An increasing body of evidence in the past 2 decades has corroborated the reciprocal relationship between pain and sleep.^[9–11] Pain and analgesic use has been shown to disturb sleep^[12,13]; on the other hand, sleep disturbance also induces hyperalgesia.^[13,14] Impaired sleep has been shown to predict new pain incidents and aggravation of chronic pain; in addition, sleep impairment is a more consistent predictor of pain than pain is of sleep.^[15–18] Experimental disturbance of sleep such as full sleep deprivation, partial sleep deprivation,^[19,20] and sleep fragmentation^[21] have all been shown to increase both clinical pain and responses to sensory tests; however, there is considerable interindividual variability in this respect.^[22] Nonetheless, the link between sleep quality and the intensity of acute postsurgical pain is not well characterized. One study used Actiwatch for evaluation of sleep on just the night before breast-conserving surgery; they found a significant negative relationship between

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sleep efficiency on the night before surgery and the severity of postoperative pain.^[23] However, assessment of sleep only on the night before operation is not enough to reflect the overall sleep status of patients.

As we know, perioperative sleep disturbance is a common problem that is frequently neglected by clinicians. In recent years, postoperative sleep disturbance has evoked much attention.^[24–26] But it is evident that many patients have already been suffering from poor sleep quality for long before surgery; this phenomenon is especially seen among breast cancer patients because most breast cancer manifests in mid life.^[27] Approximately 30% to 75% of newly diagnosed or recently treated breast cancer patients reportedly suffer from chronic poor sleep quality.^[28] Thus, examining the effect of chronically preoperative sleep disturbance on acute postoperative pain in breast cancer patients receiving total mastectomy surgery is a key imperative. This may help improve the management of acute postoperative pain, which is important to decrease the incidence of persistent chronic pain after mastectomy surgery. Therefore, we designed this prospective cohort study to investigate the relationship between sleep quality of 1 month before surgery and acute postoperative pain in women scheduled for modified radical mastectomy. We assumed that poor sleep quality before surgery will increase the intensity of acute postoperative pain and postoperative analgesic consumption.

2. Methods

2.1. Study design and setting

This study was registered before patient enrollment in the Chinese Clinical Trial Registry (ID ChiCTR-IPC-17012922). The study was conducted at the Liao-cheng People's Hospital between April 2017 and June 2018. Ethics approval was provided by the ethical committee of Liao-cheng people's hospital. Written informed consent was obtained from all participants prior to their enrolment.

2.2. Subjects

Inclusion criteria: women aged 18 to 65 years scheduled to undergo unilateral modified radical mastectomy; American Society of Anesthesiologists physical status classification I or II.

Exclusion criteria: previous breast or axillary surgery, pregnant women, women affected by anxiety or depression disorders (Self-rating Anxiety Scale score ≥ 50 or Self-rating Depression Scale score ≥ 53 , respectively), history of chronic pain including different type of headache, sleep apnea syndrome, psychiatric disorder, detectable metastatic diseases, and cardiac diseases, and inability to complete Chinese test questionnaires.

2.3. Preoperative questionnaires

A researcher, who was not involved in the treatment and follow-up of patients, performed preoperative interview on the day prior to surgery to collect demographic information and submitted questionnaires of sleep quality and mental state to the enrolled patients; all participants completed the questionnaires preoperatively.

2.3.1. Sleep quality evaluation. Preoperative subjective sleep quality: The Pittsburgh Sleep Quality Index (PSQI) questionnaire was used to evaluate the participants' subjective sleep quality of

the past month. PSQI is a standard instrument with a high reliability and validity to assess the sleep quality of patients over a 1-month period,^[29] which consists of 19 individual items grouped into 7 subscale scores: sleep duration; sleep latency (time needed to fall asleep); subjective sleep quality index; habitual sleep efficiency (proportion between total sleep time and time in bed); sleep disturbances (waking up during the night); use of sedatives; and daytime dysfunction. Each of these subscales is weighted equally on a 4-point scale generating a total score ranging from 0 to 21. Total score ≥ 5 is generally considered indicative of poor sleep quality; higher scores reflect a greater decline in sleep quality. In this study, preoperative sleep quality during the past month was assessed using the PSQI Chinese version on the day of surgery. Patients whose PSQI score were ≥ 5 were allocated to the poor sleep group, while those with PSQI score < 5 were assigned to the good sleep group.

2.3.2. Preoperative mental state. Preoperative mental state was evaluated on the day of surgery using the Chinese-language versions of the Self-rating Anxiety Scale (SAS) and Self-rating Depression Scale (SDS). SAS and SDS are self-report questionnaires for a comprehensive assessment of anxiety and depression status of patients.^[30,31] The SAS and the SDS comprised of 20 items. Each item was scored on a 4-point scale (0 – not at all; 1 – somewhat; 3 – moderately so; and 4 – very much so); the total score for each questionnaire was multiplied by 1.25 to convert it into a standardized score (range, 25–100; higher scores indicate higher levels of anxiety and depression). According to Zung severity levels classification criteria, the SDS score greater than 53 was used by Chinese psychiatric professionals as a cutoff point for depression-related symptom severity, and the cutoff values have been widely used in Chinese depression studies.^[32,33] Patients with SAS score ≥ 50 were considered to suffer from anxiety, and patients with SDS score ≥ 53 were considered to suffer from depression in this study. Patients with anxiety and depression were excluded from this study.

2.4. Anesthesia and postoperative analgesia

All enrolled patients were monitored with pulse oximetry (SPO₂), electrocardiography, noninvasive blood pressure, acceleromyography, and bispectral index monitor (BIS LoC2 Channel; COVIDINE IIc, Mansfield, MA). General anesthesia was induced for all patients with propofol 2 to 3 mg/kg and fentanyl 2 μ g/kg with no premedication. Endo-tracheal intubation was facilitated by 0.15 mg/kg cisatracurium IV; 7-mm tracheal tube was inserted by visual laryngoscopy. Anesthesia was maintained with an intravenous infusion of propofol (6–8 mg/kg/hour) and remifentanyl (0.05–0.1 μ g/kg/minute) to maintain the bispectral index within the range of 40 to 60. The presence of hypertension or tachycardia ($>20\%$ of baseline) during anesthesia was attributed to insufficient analgesia and fentanyl 1 μ g/kg was administered while BIS was 40 to 60. Cisatracurium 0.03 mg/kg was administered intermittently at a train-of-four count of ≥ 1 . Mechanical ventilation was maintained with a tidal volume of 8 mL/kg, and the ventilatory frequency was adjusted to maintain a normal end-tidal carbon dioxide concentration with an air/oxygen mixture (fraction of inspired oxygen, 0.5). Intraoperatively, Ephedrine was administered when blood pressure drops to 20% of the baseline and atropine was administered when heart rate drops to 20% of the baseline. The mid-esophageal temperature was maintained at 36 to 37°C during surgery.

Ketorolac 30 mg was injected intravenously 30 minutes before the end of the operation and neostigmine 0.05 mg/kg plus atropine 0.02 mg/kg were used to reverse possible residual neuromuscular blockade at the end of surgery. All patients were administered ondansetron 8 mg in order to prevent postoperative nausea and vomiting (PONV). The tracheal tube was removed when the patients regained their consciousness and were able to breathe spontaneously; subsequently, the patients were transferred to the postanesthesia care unit (PACU).

In the PACU, severity of pain was assessed using the numerical rating score (NRS). Fentanyl (50 μ g) was administered if the NRS score was >4 points or if the patients demanded analgesics. All participants were discharged from the PACU to the ward when the Steward Recovery Score exceeded 4 points and NRS was less than 4.

2.5. Follow-up

In the ward, all participants were followed up by the acute pain service (APS) team blinded to group allocation. NRS was used to evaluate postoperative peak pain at 0 (T0), 2 (T1), 6 (T2), 12 (T3), and 24 (T4) hours after surgery. In this study, NRS score ≥ 7 was defined as severe pain. NRS score ≥ 4 (effect on general activity, mood, and postoperative mobility)^[34] was considered indicative of clinically relevant moderate pain, which is comparable with other literature.^[35,36] Patients were encouraged to request analgesic medication at any time. Analgesics were administered once the NRS exceeded 4 or on the patient's request for analgesia. Ibuprofen 400 mg (oral) was the first choice. Nefopam 20 mg (intramuscular injection) was the rescue medication, if Ibuprofen did not help. If these NSAIDs were still inadequate, 2 mg morphine was injected intravenously.

2.6. Main outcome

The incidence of severe pain within the 24-hour postoperative period was the primary endpoint. Secondary endpoints included the maximum NRS score, the number of patients who used analgesic medication, and the number of patients who required rescued analgesic medication within the same time.

2.7. Postoperative complications and length of hospital stay

PONV, fever, and the length of hospital stay (from the day of surgery up to discharge) were also collected.

2.8. Sample size and statistical analysis

In our preliminary analysis of data from 60 patients (30 patients with good sleep quality and 30 patients with poor sleep quality), the incidence rate of severe postoperative pain was 7% and 30%, respectively. A sample size of 88 (44 in each group) was required to detect a between-group difference, with anticipated α error (2 tailed) of 0.05 and β error 0.20 (power: 0.8). Finally, a sample size of 106 (53 in each group) was required to account for 20% attrition.

All statistical analyses were performed using IBM SPSS for Windows version 22.0 (SPSS, Chicago, IL). Normality of data distribution was assessed using the Kolmogorov–Smirnov test. Demographic data are presented as mean (standard deviation [SD]) or median (interquartile range); categorical variables are presented as frequencies (percentages). Between-group differences with respect to normally distributed continuous variables were assessed using the Student *t* test, while those with respect to nonnormally distributed continuous variables were assessed using the Mann–Whitney *U* test. Fisher exact Chi-squared test, Pearson Chi-squared test, or Cochran–Mantel–Haenszel test were used to assess between-group differences with respect to categorical variables, as appropriate. A linear mixed-effects model was used to analyze the change of NRS score during the 24-hour postoperative period. A 2-sided *P* value $< .05$ was considered indicative of statistically significant difference.

3. Results

Between April 2017 and June 2018, 232 female patients were assessed for eligibility and 108 patients were enrolled in this study. Among the 108 women, 53 patients (49%) had good sleep quality while 55 (51%) patients had bad sleep quality. Three dropped out of the study during follow-up in each group. Finally, a total of 102 patients were included in the final analysis (Fig. 1). The average PSQI score was 3.4 ± 1.3 in the good sleep group and

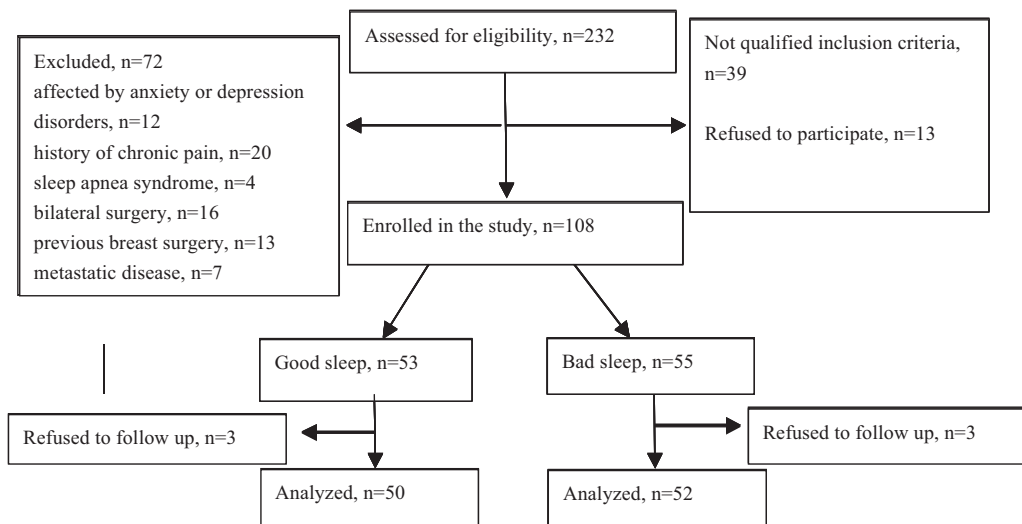


Figure 1. Schematic illustration of the study design and the patient-selection criteria.

Table 1
Demographic characteristics, preoperative mental scores, and surgical and anesthetic parameters.

	Good sleepers N=50	Poor sleepers N=52	P value
Age, y	46.80 ± 5.70	47.90 ± 5.49	.322 [*]
Height, kg	159.04 ± 4.06	157.92 ± 2.581	.099 [*]
Weight, cm	63.41 ± 6.99	62.71 ± 5.55	.577 [*]
BMI, kg/cm ²	25.07 ± 2.59	25.13 ± 2.02	.89 [*]
Single/married	4/46	4/48	.954 [†]
SAS	46 (38, 49)	47 (39, 49)	.2 [‡]
SDA	40 (35, 47)	41 (36, 49)	.11 [‡]
ASA VIII	29/21	25/27	.315 [§]
Duration of anesthesia, min	110.40 ± 17.17	112.88 ± 11.98	.397 [*]
Duration of surgery, min	96.34 ± 13.05	99.13 ± 11.58	.255 [*]
Propofol consumption, mg	815.10 ± 143.15	826.44 ± 129.77	.676 [*]
Fentanyl consumption, mg	0.2 (0.2, 0.215)	0.2 (0.2, 0.25)	.644 [‡]
Remifentanyl consumption, mg	0.58 ± 0.11	0.59 ± 0.11	.707 [*]
Cisatracurium consumption, mg	14 (14, 16)	14 (14, 16)	.377 [‡]
Blood loss, mL	92 ± 16	95 ± 12	.335 [*]
Fluid transfusion, mL	851 ± 126	894 ± 104	.061 [*]
patients need ephedrine yes/no	10/40	14/38	.410 [§]
Number of patients need atropine	8/42	12/40	.368 [§]

Data presented as mean ± SD, frequency (percentage), or as median (interquartile range). ASA = American Society of Anesthesiologists, BMI = body mass index, SAS = Self-rating Anxiety Scale, SD = standard deviation, SDS = Self-rating Depression Scale.

^{*} Independent-sample *t* test.

[†] Fisher exact test.

[‡] Mann-Whitney *U* test.

[§] Pearson Chi-squared test.

11.1 ± 2.6 in the poor sleep group. There was no significant between-group difference with respect to baseline demographic characteristics including age, height, weight, BMI, marital status, and the scores of SAS and SAD (Table 1).

There was no significant between-group difference with respect to the American Society of Anesthesiologists grade, type of surgery, duration of surgery, duration of anesthesia, blood loss, fluid transfusion, and the consumption of propofol, fentanyl, remifentanyl, cisatracurium, and vasoactive agent (Table 1).

NRS score at different time-points during the first 24 hours is shown in Table 2. The pain scores were comparable in the 2 groups at T0 ($P = .589$); patients in the poor sleep group had significantly higher postoperative pain scores than those in the good sleep group at T1 ($P = .002$), T2 ($P < .001$), T3 ($P < .001$), and T4 ($P = .002$). A significant group effect was identified with a negative coefficient (estimate = -0.87500, $P = .001$), which indi-

Table 2
Postoperative NRS score at different time-points during the first postoperative 24 hours.

Time	Good sleepers N=50	Poor sleepers N=52	P value
NRS T0	0 (0, 1)	0 (0, 1)	.589 [*]
NRS T1	1.5 (1, 2)	2 (2, 2.75)	.002 [*]
NRS T2	2 (2, 3)	3 (2, 5)	.000 [*]
NRS T3	3 (3, 4)	4.5 (3.25, 6.0)	.000 [*]
NRS T4	3 (3, 4)	4.0 (3.25, 6.0)	.002 [*]

Data presented as median (interquartile range).

NRS = numerical rating score (0–10) for pain.

^{*} Mann-Whitney *U* test.

Table 3
Effect of group and postoperative time on the change of NRS score.

	Estimate	95% CI	t value	P value
Intercept	2.7261	2.3829–3.0693	15.673	.000
Group (good sleep)	-0.8750	-1.3652–-0.3848	-3.522	.001
Time	0.08659	0.0693–0.1039	9.915	.000
Group * time	0.00118	-0.0235–0.0259	0.095	.925

CI = confidence interval, NRS = numerical rating score (0–10) for pain.

cated that the postoperative NRS scores in the good sleep group were significantly lower than those in the poor sleep group. Conversely, a significant effect of the postoperative time was identified with a positive coefficient (estimate = 0.08659, $P < .001$), which indicated that the NRS score increased within the first 24 hours postsurgery. Additionally, no significant interaction between groups and the postoperative time was found (estimate = 0.00118, $P = .925$), which indicated that the change in NRS scores was comparable in the 2 groups (Table 3).

The incidence of severe pain in the poor sleep group was higher than that in the good sleep group (8% vs 27%, $P = .018$) (Table 4). In addition, maximum NRS pain score during the first postoperative 24 hours in the poor sleep group was higher than that in the good sleep group (5 [3.25, 7] vs 3 [3, 4], $P < .001$). The percentage of patients who used analgesic medicine within the 24-hour postsurgical period in the good sleep group and the poor sleep group was 76% and 84.62%, respectively; there was no significant between-group difference in this respect ($P = .273$). However, the ratio of participants who required rescued analgesics within the same time was significantly different, 52% of participants used rescued analgesics-nefopan in the poor sleep group, compared with 22% in the good sleep group ($P = .002$); in addition, 23% of participants with poor sleep quality used rescued analgesics-morphine, compared with 6% in the good sleep group ($P = .0223$).

Data pertaining to postoperative complications and hospital stay are presented in Table 5. The incidence of PONV ($P = .035$) and postoperative fever ($P = .004$) in the poor sleep group during the first 24 hours was significantly higher than that in the good sleep group. In addition, patients with poor sleep quality had significantly longer hospital stay ($P < .001$).

Table 4
Maximum pain score, incidence of severe pain, and characterization of analgesic use during the first 24 h.

	Good sleepers N=50	Poor sleepers N=52	P value
Maximum NRS score	3 (3, 4)	5 (3.25, 7)	<.001 [*]
Number of patients experienced severe pain	4 (8%)	14 (27%)	=.018 [†]
Number of patients received analgesics	38 (76%)	44 (85%)	=.273 [‡]
Number of patients received nefopan	11 (22%)	27 (52%)	=.002 [‡]
Number of patients received morphine	3 (6%)	12 (23%)	=.0223 [†]

Data presented as median (inter-quartile range) or as frequency (percentage). NRS = numerical rating score (0–10) for pain.

^{*} Mann-Whitney *U* test.

[†] Fisher exact test.

[‡] Pearson Chi-square test.

Table 5
Postoperative complications and length of hospital stay in the 2 groups.

	Good sleepers N = 50	Poor sleepers N = 52	P value
Nausea and vomiting	11(22%)	22(42%)	=.035*
Fever	7(14%)	21(40%)	=.004*
Hospital stay, d	7.7 ± 1.2	8.7 ± 1.3	<.001†

Data presented as frequency (percentage) or mean ± standard deviation (SD).

* Pearson Chi-squared test.

† Independent-sample *t* test.

4. Discussion

As anticipated, we found that breast cancer patients with poor sleep quality before surgery had higher incidence of severe postoperative pain, reported higher NRS scores, and demanded more rescued analgesics in the first postoperative 24 hours. What is unexpected was that the incidence of PONV and postoperative fever is higher in poor sleep group; in addition, patients with poor preoperative sleep had longer hospital stay.

In this study population, 51% of breast cancer patients scheduled for breast surgery were affected by poor sleep before surgery, which is consistent with the results of previous studies.^[37,38] Adult women of all ages with breast cancer are affected by sleep disturbances. Poor sleep quality, frequent nocturnal awakenings, and insomnia are the main characteristics of sleep disorders experienced by breast cancer patients aged ≥50 years.^[37] In a previous study, 53.5% of women with no preoperative breast pain reported sleep problems before breast cancer surgery as against 66.7% of patients who had preoperative breast pain.^[38]

Of note, we found that patients with poor sleep quality before surgery were more likely to report higher postoperative pain scores (NRS); in addition, the incidence rate of severe pain was also higher in the poor sleep quality group. Although there was no significant between-group difference with respect to the ratio of participants received analgesics during the first postoperative 24 hours, the number of patients who required rescued analgesics was far greater in the poor sleep group. Our findings add to the growing body of evidence that impaired preoperative sleep is a consistent predictor of postoperative pain. In 2017, one feasibility study found that a prophylactic extended time in bed before joint replacement surgery in patients with moderately short sleep resulted in increased sleep time and reduced postoperative pain and analgesic use^[39]; this provided an indirect evidence of the relationship of preoperative sleep with postoperative pain. There were also some direct evidences. In a cross-sectional study of adults undergoing scheduled abdominal or orthopedic surgery, self-reported chronic sleep disturbance was the only predictor of severe postoperative pain at rest; it was associated with a 2- to 3-fold higher risk of severe postoperative pain.^[40] However, the authors did not use standard sleep assessment tools for a comprehensive assessment of preoperative sleep quality. Caroline et al used Actiwatch to evaluate patients' sleep on just the night before breast-conserving surgery; they found actigraphy-determined total time in bed, sleep onset time, sleep end time, and total sleep time (sleep duration) the night before surgery were not associated with postoperative pain, but there existed a significant negative relationship between sleep efficiency on the night before surgery and the severity of

postoperative pain.^[23] However, assessment of sleep only on the night before operation is not enough to reflect the overall sleep status of patients; in addition, the use of Actiwatch is liable to disrupt the patient's normal sleep and is unpractical to assess all patients' sleep condition. In clinical settings, sleep disturbance lasting for a duration of 2 to 3 weeks is typically used to differentiate between transient and persistent sleep disorders. Therefore, the use of PSQI in our study allowed us to distinguish between most transient and persistent sleep disturbances. Beyond that, PSQI is a convenient, practical, and economic tool to assess patients' sleep condition. In addition, the brightest spot of our study is that we excluded all patients with anxiety and depression, which is an important risk factor of postoperative pain. Indeed, the anxiety and depression scores of participants in the poor sleep group were slightly higher than that in the good sleep group in our study, but there was no difference between groups. Studies suggest that sleep disturbances may impair important physiological processes such as dopaminergic signaling, opioidergic signaling, and emotional regulation, which contribute to the development of hyperalgesia and maintenance of chronic pain.^[15] Since the mechanism of acute postoperative pain is different from that of chronic pain, future studies investigating the association of sleep and acute postoperative pain should also focus on the underlying mechanisms.

To the best of our knowledge, this is the first study that reveals that women with poor preoperative sleep quality are more likely to experience PONV. The incidence of PONV in the poor sleep group was almost twice as high as that in the good sleep group (42% vs 22%, $P < .05$). Indeed, women undergoing breast surgery under general anesthesia are at a particularly high risk of PONV; among patients who received no antiemetic medication, the reported incidence is 60% to 80%.^[41-43] In this study, all participants received antiemetic medication at the completion of surgery, which explains the low incidence rate of PONV in our study population. In a previous study of breast cancer patients, poor sleep quality was associated with an increased risk of chemotherapy-induced nausea and vomiting^[44] due to autonomic nervous system dysfunction induced by poor sleep. Although the side effect of nefopam is another factor to explain the higher incidence of PONV in the poor sleep group.^[45] In the past decades, numerous studies have identified the risk factors of PONA, such as female gender, history of vestibular diseases, use of opioids, and absence of a smoking history.^[46] It seems likely that preoperative sleep disturbance is another important correlate of PONV that should not be ignored. Future studies should investigate the relationship between preoperative sleep quality and PONV.

The incidence of postoperative fever was higher in the poor sleep group, which can be explained by the relationship between sleep and inflammatory system. The current viewpoint is that sleep disorder has an effect on the inflammatory system activation, which is different from direct stimulation of the immune system by infection or injury.^[47,48] Levels of interleukin (IL)-1 beta, IL-6, and tumor necrosis factor (TNF)-alpha were found to be elevated after acute sleep deprivation or chronic partial sleep deprivation in healthy participants.^[49-54] In addition, sleep disturbance (as assessed by validated questionnaires) was shown to be associated with increased levels of C-reactive protein and IL-6.^[55] However, the mechanism by which sleep disorder activates inflammatory pathways is still unclear. Further studies are required to confirm the association between preoperative sleep quality and the change in inflammatory system in patients undergoing surgery.

Some limitations of our study need to be acknowledged. Firstly, the sleep quality was only assessed using the PSQI; we did not evaluate the sleep patterns using more objective methods such as polysomnography or actigraphy. Secondly, it is not a randomised, controlled, blinded trial. It is a single-center, relatively small population trial, a multicenter trial is necessary to verify this result. Finally, acute postoperative pain was just reported within the postoperative 24 hours, and we did not examine the duration of postoperative pain and persistent postoperative pain after breast cancer surgery.

5. Conclusions

Breast cancer patients with poor sleep quality prior to surgery reported higher postoperative pain scores in the first postoperative 24 h, required more analgesics, experienced more complications (such as PONV), and had longer hospital stay. Further studies using more objective measures of sleep are required to validate our findings. We suggest that Chinese clinicians should pay more attention to preoperative sleep quality of breast cancer patients.

Author contributions

Conceptualization: Jin-ping Wang, Su-fen Lu, Li-na Guo, Chun-guang Ren, Zong-wang Zhang.

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Supervision: Jin-ping Wang, Su-fen Lu.

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