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Efficacy of a continuous wound infiltration system for postoperative pain management in gynecologic patients who underwent single-port access laparoscopy for adnexal disease

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Introduction: Single-port access (SPA) laparoscopy requires only one incision, unlike conventional laparoscopy. However, its umbilical incision is larger than that of conventional laparoscopy and can be vulnerable to postoperative pain. This study aimed to evaluate whether simultaneous use of a continuous wound infiltration (CWI) system and intravenous patient-controlled analgesia (IV PCA) effectively decreases surgical site pain in patients who underwent SPA laparoscopy due to gynecologic adnexal disease.

Methods: A total of 371 patients who underwent SPA laparoscopy and who received IV PCA or CWI was retrospectively reviewed (combined group [CWI + IV PCA, n = 159] vs. PCA group [IV PCA only, n = 212]). To evaluate postoperative pain management, the numeric rating scale (NRS) pain score after surgery, total amount of fentanyl administered via IV PCA, and additional pain killer consumption were collected.

Results: The NRS scores at 12 h (1.90 \pm 1.11 vs. 2.70 \pm 1.08, p < 0.001) and 24 h (1.82 \pm 0.82 vs. 2.11 \pm 1.44, p = 0.026) after surgery were significantly lower in the combined group than in the PCA group. The total amount of PCA fentanyl was significantly smaller in the combined group than in the PCA group (p < 0.001). The total quantity of rescue analgesics was smaller in the combined group than in the PCA group (p < 0.05).

Conclusion: Combined use of the CWI system and IV PCA is an effective postoperative pain management strategy in patient who underwent SPA laparoscopy for adnexal disease.

KEYWORDS

single-port access laparoscopy, adnexal disease, postoperative pain, continuous wound infiltration, intravenous patient-controlled analgesia

1. Introduction

Single-port access (SPA) laparoscopy has been widely performed in gynecologic disease due to its advantages over conventional laparoscopy (1, 2). SPA requires only one abdominal incision at the umbilicus, unlike conventional laparoscopy, which requires at least two or three incisions. However, the umbilical incision of SPA laparoscopy is larger than that of conventional laparoscopy because insertion of all laparoscopic instruments and specimen retrieval are performed through the umbilicus (3). Therefore, patients treated with SPA laparoscopy can be vulnerable to postoperative pain despite receiving the smallest number of abdominal incisions among minimally invasive surgeries (MISs).

Postoperative pain is one of the major concerns for patients before undergoing surgery (4). Inadequately managed postoperative pain can cause various complications such as impairment of physical function and lengthened hospital stay, increasing socioeconomic costs (5). Therefore, pain management is considered one of the key elements in the Enhanced Recovery after Surgery (ERAS) protocol (6, 7). Various pain control methods such as systemic and regional analgesics have been developed (8). Traditionally, a combination of systemic analgesics based on intravenous patient-controlled analgesia (IV PCA) has been preferred for MIS as a multimodal approach. However, the side effects associated with opioids are a disadvantage of IV PCA (9). The continuous wound infiltration (CWI) system, in which a local anesthetic is continuously applied into the surgical wound, has been demonstrated to have postoperative pain-reducing effects and opioidsparing effects in various laparotomic wounds (10, 11). However, no study has evaluated the effectiveness of a CWI system to manage postoperative pain for SPA laparoscopy for gynecologic disease, which requires the largest incision among MISs.

Therefore, the purpose of this study was to evaluate whether simultaneous use of the CWI system and IV PCA effectively decreases postoperative surgical site pain compared with IV PCA alone in patients who underwent SPA laparoscopy due to gynecologic adnexal diseases.

2. Materials and methods

2.1. Patient selection

We retrospectively reviewed patients who underwent SPA laparoscopy for adnexal disease in Kangbuk Samsung Hospital and or Uijeongbu Eulji Medical Center between January 1, 2020 and August 31, 2022. This study was approved by the Institutional Review Boards (IRBs) of Kangbuk Samsung Hospital (IRB no. 2022–12-030) and Eulji University (IRB no. 2023-02-015). Patients who met the following criteria were eligible for this study: (1) those who underwent adnexal surgery including ovarian cystectomy or salpingo-oophorectomy, (2) those who underwent IV PCA or CWI for postoperative pain control, and (3) those whose pain score at 6, 12, 24, and 48 h after surgery were adequately assessed using the numeric rating scale (NRS). We excluded patients who met one or more of the following criteria: (1) those who underwent another gynecologic surgery (such as hysterectomy or myomectomy) simultaneously, (2) those who were converted to multiport laparoscopy or laparotomy during surgery, and (3) those in which the IV PCA or CWI system was not sufficiently maintained during hospitalization because of side effects. Patients were then classified into the combined group (IV PCA and CWI system) and the PCA group (IV PCA only).

2.1.1. Data collection and definition

Patient baseline characteristics including age, body mass index (BMI), and history of abdominal surgery were collected. Total operative time, perioperative complications, estimated blood loss (EBL), serum hemoglobin (Hb) level difference, single-port placement time, specimen retrieval time, length of umbilical incision before and after surgery, duration of hospital stay, and pathologic diagnosis were recorded for surgical outcomes. The total operative time was defined as the time from skin incision to skin closure. EBL was calculated by the anesthesiology unit as the difference between the total amounts of suction and irrigation. Serum Hb level difference was defined as the change between preoperative Hb level and Hb level on postoperative day 1. The single-port placement time was defined as the time for which pneumoperitoneum was maintained by insufflating carbon dioxide (CO2) gas. We defined specimen retrieval time as the total time required to remove the resected tissue through the in-bag removal process. To evaluate umbilical incision enlargement due to the elastic wound retractor of the single-port system and specimen retrieval process during surgery, the longest vertical length of umbilical incision was measured before and after surgery with the wound retractor inserted (Supplementary Figure S1). The length of hospital stay was defined as the time from operation day to discharge day.

For postoperative pain outcomes, the NRS, ranging from 0 (no pain) to 10 (worst pain), was used to assess surgical site pain at 6, 12, 24, and 48 h after surgery. The total amount of fentanyl citrate administered via IV PCA during the entire hospitalization period was recorded. The timing and amount of additional pain killers such as nonsteroidal anti-inflammatory drugs (NSAIDs) or opioid drugs were also recorded. In both institutions, the routine pain management protocol after SPA laparoscopy for adnexal disease is as follows. All patients underwent IV PCA routinely at the postanesthesia care unit as a baseline pain management method immediately after surgery. The CWI system was performed in some patients, though there were no definite clinical criteria for its use. However, before surgery, detailed counseling for the clinical benefits, risks, and cost of the CWI system was offered to patients. The patients made their own decision whether receive the CWI system in addition to baseline IV PCA, and the system was applied only to patients who consented to the procedure. For IV PCA, 900-1800 µg fentanyl citrate was mixed with 100 mL of 0.9% normal saline (detailed criteria for the dosage of fentanyl citrate according to age and body weight are described in Supplementary Table S1); it was continuously infused at a rate of 1 cc/h. When patients wanted additional infusion for pain relief, a bolus dose of 15 μg fentanyl citrate was administered with 15-min lockout time. After starting an oral diet, oral NSAIDs were administered twice a day until discharge; in the case of allergic reaction to NSAIDs, oral acetaminophen was administered three times a day. For additional pain management, NSAIDs were used first, and opioid drugs were used when necessary. Additional pain killers were administered at any time upon patient request. Patients who underwent adnexal laparoscopic surgery were usually discharged one or two days after surgery if there were no complications.

2.1.2. Surgical procedure

The surgical technique of SPA laparoscopy has been described previously (12). Briefly, a small vertical transumbilical incision of 2.0to 2.5-cm was made using the open Hasson technique. A single-port elastic wound retractor was inserted through the umbilicus (Supplementary Figures S1A,B), and the longest vertical length of umbilical incision was measured (Supplementary Figure S1C). A single multichannel cap was placed on the wound retractor (Supplementary Figure S1D). Pneumoperitoneum with CO2 at 12 mmHg was established, and laparoscopic surgery (ovarian cystectomy or salpingo-oophorectomy) was performed. The resected adnexal tissues were placed into an endo-pouch specimen retrieval bag and removed through the umbilical opening using the cold knifein-bag tissue removal technique (Supplementary Figure S1E). At the end of the laparoscopic surgery, the length of the umbilical incision was measured again with the wound retractor inserted to evaluate enlargement of the incision (Supplementary Figure S1F). Then, the musculo-fascial layer was closed.

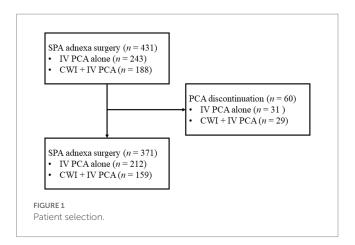
In patients receiving CWI (ON-Q PainBuster®, I-Flow Corporation, Halyard Health, Irvine, CA, United States), the guiding needle was inserted into the subcutaneous layer from the lower end of the vertical umbilical incision to a point 7-10 cm below it (Supplementary Figures S2A,B). A multi-hole soaker catheter was then inserted under the guidance of guiding needle and placed on the musculo-facial layer along the umbilical wound (Supplementary Figure S2C). The subcutaneous layer and skin were closed, and the catheter was connected to an elastomeric pump filled with 100 mL of 0.3% ropivacaine solution (Supplementary Figure S2D). This solution was continuously infused at a rate of 2 mL/h.

2.1.3. Statistical analysis

All statistical analyzes were performed using SPSS version 25.0 (IBM SPSS Statistics for Windows, IBM Corp., Armonk, NY, United States). Normality of the data was assessed with the Shapiro–Wilk test. Data with normal distribution are presented as mean \pm standard deviation (SD), while median (interquartile range, IQR) were used for data with non-normal distribution. Frequency distributions among categorical variables for the two pain control methods were compared using the chi-square test or Fisher's exact test. Multiple linear regression analysis models were used to estimate the independent contributions of variables to postoperative pain. A value of p < 0.05 was considered statistically significant.

3. Results

During the study period, 431 patients underwent SPA laparoscopy for adnexal disease and received IV PCA or CWI for postoperative pain management. Of these, 60 patients who could not maintain IV PCA until discharge due to side effects were excluded. A total of 371 patients was enrolled in this study, comprising 212 (57.1%) patients who received only IV PCA (PCA group) and 159 (42.9%) patients who received IV PCA and CWI concurrently (combined group) (Figure 1). The patient characteristics and surgical outcomes of each group are summarized in Table 1. There was no significant difference in age, BMI, prior abdominal surgery history, length of umbilical



wound before and after surgery, specimen retrieval time, operation time, and pathologic diagnosis between the two groups.

The pain control outcomes according to pain management method are presented in Table 2. The NRS pain score declined gradually with time in both groups (Figure 2). The mean NRS scores assessed at 12h (combined group vs. PCA group, 1.90±1.11 vs. 2.70 ± 1.08 , 95% CI [0.57; 1.03], p < 0.001) and 24h (1.82 ± 0.82 vs. 2.11 ± 1.44 , 95% CI [0.03; 0.55], p = 0.026) after surgery were significantly lower in the combined group compared with PCA group. For the remaining period, the pain intensity of the combined group tended to be lower than that of the PCA group, but it did not show statistical significance. The total amount of fentanyl citrate administered via IV PCA during the entire hospitalization period was significantly less in the combined group than in the PCA group (combined group vs. PCA group, $622.1 \pm 105.3 \,\mu g$ vs. $703.1 \pm 139.1 \,\mu g$, 95% CI [56.0; 86.3], p < 0.001). The percentage of patients requiring rescue analgesics after surgery was lower in the combined group than in the PCA group, and in particular, there was a significant difference between 6 and 12h after surgery (14.5% vs. 23.1%, p = 0.037). In addition, the CWI system significantly reduced the total amount of rescue analgesics administered from 6 to 12 h after surgery (0.23 ± 0.43) ampules vs. 0.35 ± 0.48 ampules, 95% CI [0.05; 0.22], p = 0.008). The use of higher potency rescue analgesics was less frequent in the combined group than the PCA group, but the difference was not significant.

In univariate and multivariate linear regression analyzes of postoperative pain score (Table 3), postoperative wound length, operative time, single-port placement time, and pain control method were associated with pain score at postoperative 6h. A larger postoperative umbilical wound (β =2.752, p=0.001), longer total operative time (β =0.013, p<0.001), and longer single-port placement time (β =0.012, p=0.002) were related to pain score at postoperative 6h. However, there was no correlation between these factors and postoperative 12, 24, and 48 h pain scores. Simultaneous use of CWI and IV PCA was an independent postoperative pain-reducing factor during the entire hospitalization period in multivariate analysis.

The incidence of side effects in both patient groups is shown in Table 4. Nausea and vomiting were significantly less frequent in the combined group than the PCA group (combined group vs. PCA group, 15.9% vs. 25.0%, p = 0.027). Wound discharge of the umbilical incision site occurred only in the combined group maintaining CWI (21.4% vs. 0%, p < 0.001). After removal of CWI, wound complications

TABLE 1 Demographic characteristics and surgical outcomes.

Characteristics	Total (<i>n</i> =371)	PCA group (<i>n</i> =212)	Combined group (n =159)	<i>p</i> -value	
Age (years)	38.16 ± 13.81	37.78 ± 13.85	38.67 ± 13.78	0.541	
BMI (kg/m²)	23.49 ± 4.12	23.80 ± 4.22	23.80 ± 4.22 23.01 ± 3.97		
ASA classification (n,%)				0.759	
I	246 (66.3)	138 (65.1)	108 (67.9)		
II	117 (31.5)	69 (32.5)	48 (30.2)		
III	8 (2.2)	5 (2.4)	3 (1.9)		
Previous OP history (n, %)				0.241	
Laparoscopy	66 (17.7)	33 (15.5)	33 (20.7)		
Laparotomy	70 (18.8)	38 (17.9)	32 (20.1)		
Operation type (n, %)				0.076	
Cystectomy	254 (68.5)	153 (72.2)	101 (63.5)		
Adnexectomy	117 (31.5)	59 (27.8)	58 (36.5)		
EBL (mL)	50 (30–50)	50 (30-50)	50 (30–60)	0.462	
Hb change (mg/dL)	1.95 ± 0.95	1.84 ± 0.99	2.09 ± 0.89	0.010 ^a	
Pre OP wound length (cm)	2.50 ± 0.06	2.50 ± 0.06	2.50 ± 0.05	0.980	
Post OP wound length (cm)	2.56 ± 0.08	2.56 ± 0.09	2.57 ± 0.08	0.442	
OP time (min)	56.67 ± 21.32	55.19 ± 18.86	58.65±24.14	0.122	
Single-port placement time (min)	34.92 ± 19.16	33.75 ± 15.98	36.45 ± 22.64	0.184	
Specimen retrieval time (min)	1.59 ± 1.60	1.48±0.994	1.73±2.16	0.181	
Hospital stay (days)	3 (2-3)	3 (2-3)	3 (2-3)	0.699	
Diagnosis (n, %)				0.367	
Teratoma	91 (24.5)	59 (27.8)	32 (20.1)		
Endometriosis	131 (35.3)	68 (32.1)	63 (39.6)		
Other benign conditions	135 (36.4)	78 (36.8)	57 (35.8)		
Borderline ovarian tumor	9 (2.4)	5 (2.4)	4 (2.5)		
Ovarian cancer	5 (1.3)	2 (0.9)	3 (1.9)		

PCA, Patient-Controlled Aanalgesia; BMI, Body Mass Index; ASA, American Society of Anesthesiologists physical status; OP, Operation; EBL, Expected Blood Loss; Hb, Hemoglobin. Values are given as mean ± standard deviation or number (percentage).

*p < 0.05.

such as infection and hernia were not different between the two groups.

4. Discussion

This study evaluated the efficacy of the CWI system for postoperative pain management after SPA laparoscopy in patients with gynecologic adnexal disease. We found that simultaneous use of the CWI system and IV PCA (combined group) significantly reduced the immediate postoperative pain scores and the total amount of IV PCA fentanyl during the entire hospitalization. We also found that the CWI system reduces the use of additional analgesics and the incidence of adverse events such as nausea and vomiting.

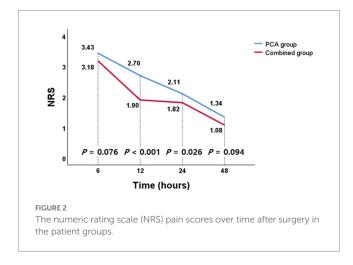
Paints with gynecologic adnexal diseases may suffer from disease-specific pain before surgery and surgery-related pain after surgery. For instance, endometriosis is a representative disease that cause sever pelvic pain by creating inflammatory change, activating immune response, and increasing innervation in the affected tissue (13–15). In addition, laparoscopic surgery itself causes various types

of pain such as surgical site pain, shoulder pain, and visceral pain (16, 17). MIS is considered the gold standard for benign gynecologic disease due to reduced postoperative pain, better cosmetic results, and faster recovery time compared with open surgery (18, 19). Among MISs, SPA laparoscopy is the least invasive in terms of number of abdominal scars but has the disadvantage of requiring the largest incision. Pain after abdominal surgery is generally known to be caused by the abdominal wall incision (20, 21). In laparoscopy, Ebanga et al. (22) reported that among surgical predictors of pain after laparoscopy, only fascia closure and increased operative time were related to immediate postoperative pain. In this respect, SPA laparoscopy has several potential risk factors that can aggravate surgical site pain. First, during the SPA laparoscopic procedure, at least two 5- to 10-mm laparoscopic instruments must be inserted simultaneously through a single umbilical opening. Collisions between instruments and non-ergonomic position of the surgeon can exert unintended forces on the umbilical incision. Second, unlike conventional laparoscopy, the umbilical wound is subjected to continuous tension from the elastic wound retractor of the singleport system while maintaining the pneumoperitoneum

TABLE 2 Pain control outcomes.

Outcomes	Total (<i>n</i> =371)	PCA group (<i>n</i> =212)	Combined group (n =159)	p-value	
NRS, mean ± SD					
Post OP 6h	3.33 ± 1.35	3.43 ± 1.43	3.18 ± 1.22	0.076	
Post OP 12 h	2.36 ± 1.16	2.70 ± 1.08	1.90 ± 1.11	<0.001ª	
Post OP 24 h	1.98 ± 1.21	2.11 ± 1.44	1.82 ± 0.82	0.026ª	
Post OP 48 h	1.22 ± 1.39	1.34 ± 1.51	1.08 ± 1.23	0.094	
PCA fentanyl quantity (μg)	669.3 ± 123.7	703.1 ± 139.1	622.1 ± 105.3	<0.001ª	
Use of additional pain killer, <i>n</i> (%)			·		
Within 6 h	126 (33.9)	80 (37.7)	46 (28.9)	0.076	
6-12 h	72 (19.4)	49 (23.1)	23 (14.5)	0.037ª	
12-24 h	38 (10.2)	25 (11.8)	13 (8.2)	0.256	
24-48 h	19 (5.1)	12 (5.7)	7 (4.4)	0.586	
Number of additional painkiller ampu	ules used, mean ± SD				
Within 6h	0.54 ± 0.44	0.56 ± 0.47	0.52 ± 0.42	0.425	
6–12 h	0.30 ± 0.46	0.35 ± 0.48	0.23 ± 0.43	0.008 ^a	
12-24 h	0.12 ± 0.31	0.14 ± 0.323	0.11 ± 0.27	0.246	
24-48 h	0.05 ± 0.22	0.06 ± 0.232	0.04 ± 0.588	0.558	
Type of additional painkiller used, n ((%)				
NSAIDs	157 (42.3)	90 (42.5)	67 (42.1)	0.952	
Tramadol hydrochloride	48 (12.9)	32 (15.1)	16 (10.1)	0.153	
Pethidine hydrochloride	27 (7.8)	20 (9.4)	7 (4.4)	0.071	
Morphine sulfate	0	0	0	-	

NRS, Numeric Rating Scale; PCA, Patient-Controlled Analgesia; OP, Operation; NSAIDs, Nonsteroidal Anti-Inflammatory Drugs. Values are given as mean \pm standard deviation or number (percentage). *p < 0.05.



(Supplementary Figure S1B). Third, the retrieval of specimens such as huge ovarian cyst or myoma is performed through the umbilicus using an in-bag tissue removal process (Supplementary Figure S1E). During this procedure, extensive wound traction for a long time can result in enlargement of the umbilical wound. Oh et al. (23) reported that the transumbilical morcellation procedure cause umbilical incision enlargement of approximately 3 mm (from 2.5 to 2.8 cm) after surgery. In our study, wound enlargement of approximately 1 mm was observed after surgery (Table 1). This small extension can

be interpreted as the fact that the adnexal tissue was relatively soft to morcellate and the specimen retrieval time was short (<2 min). The umbilical wound of SPA laparoscopy can be vulnerable to pain, for the reasons mentioned above. While previous studies have evaluated postoperative pain outcome after SPA laparoscopy, there is still no consensus on this issue (24–26). Kim et al. (26) reported that SPA surgery is more painful than conventional laparoscopy. Therefore, establishing an effective strategy to reduce pain after SPA surgery is important. Indeed, various pharmacological and technical interventions such as laparoscopic surgery under spinal anesthesia (27, 28) and bupivacaine injection to trocar sites (29) have been developed to reduce pain after laparoscopy.

The ERAS protocols to reduce hospitalization period and morbidity have become the standard for postoperative patient care after gynecologic surgery. Adequate postoperative pain management through combined use of analgesics with a different mechanism of action is a key component of the ERAS pathway. Furthermore, since the pain response to the same surgery and routine postoperative pain management for each patient, it is also important to use an appropriate rescue analgesics tailored to each patient's characteristics (30). Recently, the 2020 American Association of Gynecologic Laparoscopists (AAGL) guidelines recommended multimodal opioid-sparing analgesia for immediate pain management after MIS (31). Opioid-based IV PCA is the preferred traditional pain control method for controlling breakthrough pain in MIS as well as open surgery. However, systemic side effects

TABLE 3 Risk factors for postoperative pain according to elapsed time after surgery.

Characteristics		Univariate			Multivariate			
	β	SE	R ²	р	β	SE	R ²	p-value
Post OP. 6h								
Age	-0.09	0.005	0.008	0.088	-	_	-	
BMI	0.013	0.017	0.002	0.444	-	-	-	
Post OP wound length	2.752	0.792	0.032	0.001ª	-	-	-	
OP time	0.013	0.003	0.039	<0.001a	0.013	0.003	0.051	<0.001ª
Single-port time	0.012	0.004	0.027	0.002ª	-	-	-	
Specimen retrieval time	0.052	0.044	0.004	0.242	-	-	-	
Pain control method ^b	-0.252	0.142	0.008	0.076	-0.306	0.142	0.051	0.032ª
Post OP. 12h			I	I				
Age	-0.005	0.004	0.003	0.280	-	_	_	_
BMI	0.003	0.015	0.000	0.855	_	_	_	_
Post OP wound length	0.240	0.693	0.000	0.729	_	_	_	_
OP time	0.001	0.003	0.000	0.790	-	-	_	_
Single-port time	-0.001	0.003	0.001	0.656	_	-	_	_
Specimen retrieval time	-0.025	0.038	0.001	0.506	-	-	_	_
Pain control method ^b	-0.796	0.116	0.115	<0.001a	-0.812	0.119	0.118	<0.001a
Post OP. 24h			I	I				
Age	-0.009	0.005	0.011	0.047ª	-	_	_	_
BMI	-0.019	0.016	0.004	0.234	-	-	-	-
Post OP wound length	-0.251	0.754	0.000	0.739	-	-	_	_
OP time	0.001	0.003	0.000	0.864	-	_	_	_
Single-port time	0.001	0.003	0.000	0.827	_	-	_	_
Specimen retrieval time	-0.005	0.041	0.000	0.904	_	_	_	_
Pain control method ^b	-0.291	0.131	0.014	0.026ª	-0.297	0.135	0.011	0.028ª
Post OP. 48 h								
Age	-0.013	0.006	0.015	0.026ª	_	_	_	_
BMI	-0.011	0.018	0.001	0.554	-	_	_	_
Post OP wound length	0.342	0.895	0.000	0.703	-	_	_	_
OP time	0.005	0.004	0.005	0.183	-	_	_	_
Single-port time	0.005	0.004	0.004	0.279	_	_	_	_
Specimen retrieval time	0.031	0.047	0.001	0.520	-	-	_	_
Pain control method ^b	-0.259	0.154	0.009	0.094	-0.012	0.006	0.010	0.045ª

BMI, Body Mass Index; OP, Operation, R^2 , Coefficient of Determination; β , Regression Coefficient; SE, Standard Error. P < 0.05. P = 100 Pain Control Analgesia alone is the reference category.

related to opioids such as nausea, vomiting, ileus, and respiratory depression are frequently reported, (32) and socio-economic costs increase due to delayed recovery and medication to alleviate these side effects. Jung et al. (33) reported that although fentanyl is an opioid with relatively few side effects, approximately 24% of patients who receive fentanyl-based IV PCA experience nausea and vomiting. In addition, according to one meta-analysis (34), opioid-based epidural PCA does not reduce recovery time compared with ERAS pain control strategy. Choi et al. (35) suggested that opioid-based IV PCA is not an essential pain control method for patients who underwent MIS for colorectal cancer. Therefore, opioid-based PCA may be inconsistent with the main purpose of the ERAS

pathway, which is to reduce opioid consumption and accelerate recovery. In our study, 60 patients discontinued IV PCA due to opioid-related systemic side effects, and even in patients who maintained IV PCA until discharge, approximately 21% complained of nausea and vomiting and received additional intervention such as antiemetics. Therefore, there is need to establish an effective strategy to reduce opioid usage and postoperative pain, especially for small wounds.

The CWI system effectively controls pain by directly injecting a local anesthetic agent into the surgical site. The ERAS group also recommended that incisional infiltration has no systemic side effects when used appropriately, and it should be incorporated into other

TABLE 4 Side effects in patient groups.

Variable	Total (n=371)	PCA group (n=212)	Combined group (n=159)	<i>p</i> -value		
Nausea/vomiting (n, %)	77 (20.7)	53 (25.0)	24 (15.9)	0.027ª		
Dizziness (n, %)	6 (1.6)	2 (0.9)	4 (2.5%)	0.409		
Hypotension (n, %)	4 (1.1)	1 (0.5)	3 (1.9)	0.318		
Wound complication $(n, \%)$						
Wound infection	10 (2.6)	7(3.3)	4 (2.5)	0.764		
Wound discharge	34 (9.2)	0	34 (21.4)	<0.001ª		
Hernia	3 (0.8)	2 (0.9)	1 (0.6)	1.000		

PCA, Patient-Controlled Analgesia. Values are given as number (percentage) *p < 0.05.

pain control methods as a component of multimodal analgesia (6). Many previous studies have demonstrated the effectiveness of the CWI system for postoperative pain control (10, 11, 36). In particular, systematic review and meta-analysis studies reported that CWI is safe and has postoperative pain reduction and opioid-sparing effects in cesarean section (37-39). However, most of these studies have focused on the large laparotomic wound and not the small laparoscopic wound. Lee et al. (40) reported that wound infiltration with short-acting local anesthetic agents effectively reduced immediate postoperative pain in the SPA laparoscopic wound. However, this one-shot administration method has a limitation in that long-term pain control is difficult due to the short half-life of the local anesthetic agent. The CWI system can be a very attractive method because it can continuously deliver a local anesthetic agent. Only a few studies have been conducted to evaluate the effectiveness of the CWI system for MIS. Oh et al. (41) evaluated the efficacy of the ropivacaine-based CWI system after laparoscopic colorectal surgery and demonstrated that it significantly reduced postoperative pain and has opioid-sparing effects. There has been no study on application of a CWI system to gynecologic SPA laparoscopy, to the best of our knowledge. In our study, simultaneous use of the CWI system and IV PCA (combined group) significantly reduced immediate postoperative pain, total amount of additional analgesia, and total quantity of opioid. In contrast, acute wound complications such as wound discharge (woozing of local anesthetic agent) have been reported only in the combined group, but there was no difference in chronic wound complications such as infection and umbilical hernia between the two groups. Furthermore, the incidence of umbilical hernia in the combined group was similar to previously reported results (0.4%) (42).

Our study demonstrated that combined use of the CWI system and IV PCA may be a safe and effective strategy to manage immediate postoperative pain in patients who underwent SPA laparoscopic adnexal surgery. Large-scale randomized trials are needed to evaluate the effectiveness of CWI in SPA laparoscopy.

To the best of our knowledge, this is the first study to evaluate the efficacy of the CWI system for gynecologic SPA laparoscopy. Another strength of this study is that it was conducted with a relatively large number of patients. However, this study has some limitations. First, this was not a prospective, randomized clinical trial. Second, among various types of pain that may occur after laparoscopic surgery such as gas-induced abdominal pain, shoulder pain, and visceral pain (16, 17), only surgical site pain was evaluated in this study. Third, there

was no comparison with a CWI system-only group. Future studies should be performed to compare PCA alone, CWI alone, and a combination strategy.

Data availability statement

The raw data supporting the conclusions of this article will be made available by the authors, without undue reservation.

Author contributions

J-HK: conceptualization, software, visualization, and writing-original draft. J-HK and KL: data curation, investigation, and project administration. J-HK and EP: formal analysis. J-HK and WK: methodology. EP and YH: validation. KL and YH: resources. EP and WK: supervision. EP, KL, and WK: writing-review and editing. All authors contributed to the article and approved the submitted version.

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

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Supplementary material

The Supplementary material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/fmed.2023.1199428/full#supplementary-material

References

- 1. Fagotti A, Bottoni C, Vizzielli G, Rossitto C, Tortorella L, Monterossi G, et al. Laparoendoscopic single-site surgery (LESS) for treatment of benign adnexal disease: single-center experience over 3-years. *J Minim Invasive Gynecol.* (2012) 19:695–700. doi: 10.1016/j.jmig.2012.07.002
- 2. Song T, Cho J, Kim TJ, Kim IR, Hahm TS, Kim BG, et al. Cosmetic outcomes of laparoendoscopic single-site hysterectomy compared with multi-port surgery: randomized controlled trial. *J Minim Invasive Gynecol.* (2013) 20:460–7. doi: 10.1016/j. jmig.2013.01.010
- 3. Kang JH, Noh JJ, Jeong SY, Shim JI, Lee YY, Choi CH, et al. Feasibility of single-port access (SPA) laparoscopy for large ovarian tumor suspected to be borderline ovarian tumor. *Front Oncol.* (2020) 10:583515. doi: 10.3389/fonc.2020.583515
- 4. Ustunel F, Erden S. Evaluation of fear of pain among surgical patients in the preoperative period. *J Perianesth Nurs.* (2022) 37:188–93. doi: 10.1016/j.jopan.2021.02.003
- 5. Sinatra R. Causes and consequences of inadequate management of acute pain. *Pain Med.* (2010) 11:1859–71. doi: 10.1111/j.1526-4637.2010.00983.x
- 6. Nelson G, Bakkum-Gamez J, Kalogera E, Glaser G, Altman A, Meyer LA, et al. Guidelines for perioperative care in gynecologic/oncology: enhanced recovery after surgery (ERAS) society recommendations-2019 update. *Int J Gynecol Cancer*. (2019) 29:651–68. doi: 10.1136/ijgc-2019-000356
- 7. Simpson JC, Bao X, Agarwala A. Pain Management in Enhanced Recovery after surgery (ERAS) protocols. *Clin Colon Rectal Surg.* (2019) 32:121–8. doi: 10.1055/s-0038-1676477
- 8. Small C, Laycock H. Acute postoperative pain management. Br J Surg. (2020) 107:e70–80. doi: 10.1002/bjs.11477
- 9. Yang Y, Wu J, Li H, Ye S, Xu X, Cheng L, et al. Prospective investigation of intravenous patient-controlled analgesia with hydromorphone or sufentanil: impact on mood, opioid adverse effects, and recovery. *BMC Anesthesiol.* (2018) 18:37. doi: 10.1186/s12871-018-0500-1
- 10. Lee B, Kim K, Ahn S, Shin HJ, Suh DH, No JH, et al. Continuous wound infiltration system for postoperative pain management in gynecologic oncology patients. *Arch Gynecol Obstet.* (2017) 295:1219–26. doi: 10.1007/s00404-017-4342-8
- 11. Chan SK, Lai PB, Li PT, Wong J, Karmakar MK, Lee KF, et al. The analgesic efficacy of continuous wound instillation with ropivacaine after open hepatic surgery. *Anaesthesia*. (2010) 65:1180–6. doi: 10.1111/j.1365-2044.2010.06530.x
- 12. Song T, Kim TJ, Lee YY, Choi CH, Lee JW, Kim BG, et al. Learning curves for single-site laparoscopic ovarian surgery. *J Minim Invasive Gynecol.* (2012) 19:344–9. doi: 10.1016/j.jmig.2012.01.003
- 13. Laganà AS, La Rosa VL, AMC R, Valenti G, Sapia F, Chiofalo B, et al. Anxiety and depression in patients with endometriosis: impact and management challenges. *Int J Women's Health*. (2017) 9:323–30. doi: 10.2147/IJWH.S119729
- 14. Scheerer C, Bauer P, Chiantera V, Sehouli J, Kaufmann A, Mechsner S. Characterization of endometriosis-associated immune cell infiltrates (EMaICI). *Arch Gynecol Obstet*. (2016) 294:657–64. doi: 10.1007/s00404-016-4142-6
- 15. Scheerer C, Frangini S, Chiantera V, Mechsner S. Reduced sympathetic innervation in endometriosis is associated to Semaphorin 3C and 3F expression. *Mol Neurobiol.* (2017) 54:5131–41. doi: 10.1007/s12035-016-0058-1
- 16. Choi JB, Kang K, Song MK, Seok S, Kim YH, Kim JE. Pain characteristics after Total laparoscopic hysterectomy. *Int J Med Sci.* (2016) 13:562–8. doi: 10.7150/ijms.15875
- 17. Li XY, Tian M, Li AZ, Han CL, Li KZ. The risk of shoulder pain after laparoscopic surgery for infertility is higher in thin patients. Sci Rep. (2021) 11:13421. doi: 10.1038/s41598-021-92762-3
- 18. Mais V, Ajossa S, Piras B, Marongiu D, Guerriero S, Melis GB. Treatment of nonendometriotic benign adnexal cysts: a randomized comparison of laparoscopy and laparotomy. *Obstet Gynecol.* (1995) 86:770–4. doi: 10.1016/0029-7844(95)00261-O
- 19. Yeung PP Jr, Bolden CR, Westreich D, Sobolewski C. Patient preferences of cosmesis for abdominal incisions in gynecologic surgery. *J Minim Invasive Gynecol.* (2013) 20:79–84. doi: 10.1016/j.jmig.2012.09.008
- 20. Pogatzki-Zahn EM, Segelcke D, Schug SA. Postoperative pain-from mechanisms to treatment. *Pain Rep.* (2017) 2:e588. doi: 10.1097/PR9.0000000000000588
- 21. Loos MJ, Scheltinga MR, Mulders LG, Roumen RM. The Pfannenstiel incision as a source of chronic pain. *Obstet Gynecol.* (2008) 111:839–46. doi: 10.1097/AOG.0b013e31816a4efa
- 22. Ebanga L, Dabi Y, Benichou J, Miailhe G, Kolanska K, Uzan J, et al. Surgical determinants of post operative pain in patients undergoing laparoscopic Adnexectomy. *J Investig Surg.* (2022) 35:1386–91. doi: 10.1080/08941939.2022.2045395
- 23. Oh SJ, Lee SY, Kim WY, Kang J, Han KH, Lee SH, et al. Comparison between transumbilical and transvaginal morcellation of a large uterus during single-port-access

- total laparoscopic hysterectomy. *Obstet Gynecol Sci.* (2020) 63:379–86. doi: 10.5468/ogs.2020.63.3.379
- 24. Kim TJ, Lee YY, An JJ, Choi CH, Lee JW, Kim BG, et al. Does single-port access (SPA) laparoscopy mean reduced pain? A retrospective cohort analysis between SPA and conventional laparoscopy. *Eur J Obstet Gynecol Reprod Biol.* (2012) 162:71–4. doi: 10.1016/j.ejogrb.2012.01.007
- 25. Schmitt A, Crochet P, Knight S, Tourette C, Loundou A, Agostini A. Single-port laparoscopy vs conventional laparoscopy in benign adnexal diseases: a systematic review and Meta-analysis. *J Minim Invasive Gynecol.* (2017) 24:1083–95. doi: 10.1016/j. jmig.2017.07.001
- 26. Kim HO, Yoo CH, Lee SR, Son BH, Park YL, Shin JH, et al. Pain after laparoscopic appendectomy: a comparison of transumbilical single-port and conventional laparoscopic surgery. *J Korean Surg Soc.* (2012) 82:172–8. doi: 10.4174/jkss.2012.82.3.172
- 27. Della Corte L, Mercorio A, Morra I, Riemma G, de Franciscis P, Palumbo M, et al. Spinal anesthesia versus general anesthesia in gynecological laparoscopic surgery: a systematic review and Meta-analysis. *Gynecol Obstet Investig.* (2022) 87:1–11. doi: 10.1159/000521364
- 28. Giampaolino P, Della Corte L, Mercorio A, Bruzzese D, Coviello A, Grasso G, et al. Laparoscopic gynecological surgery under minimally invasive anesthesia: a prospective cohort study. $Updat\ Surg.\ (2022)\ 74:1755-62.\ doi: 10.1007/s13304-022-01310-9$
- 29. Hortu I, Turkay U, Terzi H, Kale A, Yilmaz M, Balcı C, et al. Impact of bupivacaine injection to trocar sites on postoperative pain following laparoscopic hysterectomy: results from a prospective, multicentre, double-blind randomized controlled trial. *Eur J Obstet Gynecol Reprod Biol.* (2020) 252:317–22. doi: 10.1016/j.ejogrb.2020.07.007
- 30. Pinto PR, McIntyre T, Fonseca C, Almeida A, Araújo-Soares V. Pre- and post-surgical factors that predict the provision of rescue analgesia following hysterectomy. $Eur\,J\,Pain.\,(2013)\,17:423-33.\,$ doi: 10.1002/j.1532-2149.2012.00205.x
- 31. Stone R, Carey E, Fader AN, Fitzgerald J, Hammons L, Nensi A, et al. Enhanced recovery and surgical optimization protocol for minimally invasive gynecologic surgery: An AAGL white paper. *J Minim Invasive Gynecol.* (2021) 28:179–203. doi: 10.1016/j. jmig.2020.08.006
- 32. Dinges HC, Otto S, Stay DK, Bäumlein S, Waldmann S, Kranke P, et al. Side effect rates of opioids in Equianalgesic doses via intravenous patient-controlled analgesia: a systematic review and network Meta-analysis. *Anesth Analg.* (2019) 129:1153–62. doi: 10.1213/ANE.0000000000003887
- 33. Jung H, Lee KH, Jeong Y, Lee KH, Yoon S, Kim WH, et al. Effect of fentanyl-based intravenous patient-controlled analgesia with and without basal infusion on postoperative opioid consumption and opioid-related side effects: a retrospective cohort study. *J Pain Res.* (2020) 13:3095–106. doi: 10.2147/JPR.S281041
- $34.\ Hughes\ MJ,\ Ventham\ NT,\ McNally\ S,\ Harrison\ E,\ Wigmore\ S.\ Analgesia\ after\ open\ abdominal\ surgery\ in\ the\ setting\ of\ enhanced\ recovery\ surgery:\ a\ systematic\ review\ and\ meta-analysis.\ \emph{JAMA\ Surg.}\ (2014)\ 149:1224-30.\ doi:\ 10.1001/jamasurg.2014.210$
- 35. Choi YY, Park JS, Park SY, Kim HJ, Yeo J, Kim J-C, et al. Can intravenous patient-controlled analgesia be omitted in patients undergoing laparoscopic surgery for colorectal cancer? *Ann Surg Treat Res.* (2015) 88:86–91. doi: 10.4174/astr.2015.88.2.86
- 36. Wagner-Kovacec J, Povalej-Brzan P, Mekis D. Efficacy of continuous in-wound infusion of levobupivacaine and ketorolac for post-caesarean section analgesia: a prospective, randomised, double-blind, placebo-controlled trial. *BMC Anesthesiol.* (2018) 18:165. doi: 10.1186/s12871-018-0609-2
- 37. Ventham NT, Hughes M, O'Neill S, Johns N, Brady RR, Wigmore SJ. Systematic review and meta-analysis of continuous local anaesthetic wound infiltration versus epidural analgesia for postoperative pain following abdominal surgery. *Br J Surg.* (2013) 100:1280–9. doi: 10.1002/bjs.9204
- 38. Adesope O, Ituk U, Habib AS. Local anaesthetic wound infiltration for postcaesarean section analgesia: a systematic review and meta-analysis. *Eur J Anaesthesiol.* (2016) 33:731–42. doi: 10.1097/EJA.0000000000000462
- 39. Abate SM, Mergia G, Nega S, Basu B, Tadesse M. Efficacy and safety of wound infiltration modalities for postoperative pain management after cesarean section: a systematic review and network meta-analysis protocol. *Syst Rev.* (2022) 11:194. doi: 10.1186/s13643-022-02068-2
- 40. Lee JH, Cho SH, Eoh KJ, Lee JY, Nam EJ, Kim S, et al. Effect of bupivacaine versus lidocaine local anesthesia on postoperative pain reduction in single-port access laparoscopic adnexal surgery using propensity score matching. *Obstet Gynecol Sci.* (2020) 63:363–9. doi: 10.5468/ogs.2020.63.3.363
- 41. Oh BY, Park YA, Koo HY, Yun SH, Kim HC, Lee WY, et al. Analgesic efficacy of ropivacaine wound infusion after laparoscopic colorectal surgery. *Ann Surg Treat Res.* (2016) 91:202–6. doi: 10.4174/astr.2016.91.4.202
- 42. Park JY, Kim TJ, Kang HJ, Lee YY, Choi CH, Lee JW, et al. Laparoendoscopic single site (LESS) surgery in benign gynecology: perioperative and late complications of 515 cases. Eur J Obstet Gynecol Reprod Biol. (2013) 167:215–8. doi: 10.1016/j.ejogrb.2012.11.027