The Analgesic Efficacy of Ultrasound-Guided Bilateral Transversus Thoracic Muscle Plane Block After Open-Heart Surgeries: A Randomized Controlled Study

Mohamed Ahmed Hamed, Maged Labib Boules, Mina Mahrous Sobhy, Mahdy Ahmed Abdelhady

Department of Anesthesiology, Faculty of Medicine, Fayoum University, Fayoum, Egypt

Correspondence: Mohamed Ahmed Hamed, Department of Anesthesiology, Faculty of Medicine, Fayoum University, Fayoum, 63511, Egypt, Tel +20 111-856-0065, Email mah07@fayoum.edu.eg

Background: We aimed to evaluate the analgesic efficacy of ultrasound-guided bilateral transversus thoracic muscle plane block after open-heart surgeries.

Methods: Seventy patients aged above 18 years and scheduled for valve replacement or adult congenital via median sternotomy were enrolled in this study. Patients were divided into two groups, randomized by computer-generated random numbers: the block group, which had the ultrasound-guided bilateral transversus thoracic muscle plane block, and the control group, which had a sham block. The primary outcome was total fentanyl consumption in the first 24-hours. The secondary outcomes were pain score, time to the first analgesic request, time to extubation, ICU stays, and hospital stay.

Results: The total fentanyl consumption in the first 24 hours was significantly lower in the block group, with a mean difference of $-158.286$ (95% CI = (-179.271 to -137.300; $p =<0.0001$)). The time to the first analgesic request was statistically significantly shorter in the non-block group (median 3 hours) than the block group (median 14 hours). During the postoperative period (0.5–24 hours), at-rest pain scores were 1.86 units lower in the block group (the estimate was $-1.80$, 95% CI = $-2.14$ to $-1.45$, $t = -10.323$ with $p < 0.0001$). Likewise, pain scores with cough were 3.29 units lower in the block group (the estimate was $-3.29$, 95% CI = $-3.80$ to $-2.77$, $t = -12.703$, $p < 0.0001$).

Conclusion: Bilateral transversus thoracic muscle plane block is a promising and effective technique in reducing opioid consumption and controlling post-sternotomy pain after open-heart surgery via median sternotomy.

Keywords: transversus thoracic muscle plane block, fentanyl consumption, sternotomy, postoperative pain

Introduction

Postoperative pain management was dependent mainly on opioids, but the optimal results could not be achieved due to opioid-related side effects.³ The multimodal analgesic technique has emerged to avoid these adverse effects of opioids and achieve appropriate analgesia; one of its components is fascial plane blocks.⁴ Various techniques have been investigated for cardiac surgery
via median sternotomy. Including; paravertebral nerve block, the pecto intercostal fascial (PIF),\textsuperscript{15} serratus anterior, PECSI/II, erector spinae plane, and transversus thoracic muscle plane (TTMP) block.\textsuperscript{4}

TTMP block was first described by Ueshima et al\textsuperscript{5} it can block multiple anterior branches of the intercostal nerves (T 2–6),\textsuperscript{6} which dominate the internal mammary region, including the sternum.\textsuperscript{7}

We hypothesized that bilateral TTMP block would provide more effective and prolonged pain relief after sternotomy involved open-heart surgery and could be used as a multimodal opioid-sparing analgesia technique. Therefore, this study aimed to evaluate the analgesic efficacy of bilateral TTMP block after acute poststernotomy pain following cardiac surgery. The primary study outcome was the 24-hours total fentanyl consumption.

Methods
This prospective, randomized, observer-blind study was conducted following the tenets of the Declaration of Helsinki. The study design was approved by the ethical review board of Fayoum University Hospital (D 199), and written informed consent was acquired from all patients. The study was conducted after registration on ClinicalTrials.gov (NCT04116554; principal investigator: Mohamed Ahmed Hamed; date of registration: October 4, 2019, with no plan to share individual participant data). In this randomized prospective double-blinded study, seventy patients were scheduled for open-heart surgery, including valve replacement or adult congenital (ASD or VSD), in Fayoum University Hospital from December 2019 to February 2021. This study adheres to the applicable CONSORT guidelines.

Study Population
Seventy patients aged above 18 years and scheduled for elective cardiac surgery for valve replacement or adult congenital (VSD or ASD) via median sternotomy were included.

The exclusion criteria were as follows; emergency surgeries, re-do surgery, coagulopathy, preoperative poor left ventricular function (Ejection fraction (EF) <35%), systemic infections or infections at the site of injection, neuromuscular disease, psychiatric illnesses, narcotic dependency, allergy to the drug used and prolonged intensive care unit (ICU) stay for reasons as heart failure and reoperation for hemostasis.

Patients were randomly divided into two groups with 1:1 allocation (block-group and non-block group) via computer-generated random numbers placed in separate opaque envelopes and opened by study investigators just before the block. All blocks were accomplished with the same anesthesiologist. Neither patients nor the functional data collector were aware of randomization.

Anesthetic Technique
All patients were preoperatively examined and investigated by complete blood count, coagulation profile, renal functions, and electrolytes. Electrocardiography, chest x-ray, and echocardiography were routinely done. Coronary angiography and carotid arterial duplex could be requested on-demand.

Patients were premedicated by intramuscular injection of 10mg morphine on the morning of the operation. Before induction of anesthesia, a five-lead electrocardiography system was applied to monitor heart rate, rhythm, and ST segments (leads II and V\textsubscript{s}). A pulse oximeter probe was attached, and a peripheral venous cannula was placed. A 20 G cannula was inserted into either right or left radial artery under local anesthesia to measure arterial pressure and blood sampling. After pre-oxygenation, general anesthesia was induced by midazolam 2 mg, fentanyl (10\textmu g.kg\textsuperscript{-1}), propofol (3mg.Kg\textsuperscript{-1}), followed by atracurium (0.5 mg.kg\textsuperscript{-1}).

The trachea was intubated; patients were mechanically ventilated with oxygen in the air, ventilation parameters were adjusted to achieve normocarbia. An esophageal temperature probe and a urinary catheter also had been placed.

A triple-lumen central venous catheter was inserted via the right internal jugular vein for drug infusion.

Anesthesia was maintained by inhaling Isoflurane 0.4% to 1% and atracurium infusion of 0.5 mg.kg\textsuperscript{-1}.h\textsuperscript{-1} for continued muscle relaxation. During extracorporeal circulation, patients have been received propofol infusion at a rate of 50–100 \mu g.kg\textsuperscript{-1}.min\textsuperscript{-1}. In addition to atracurium infusion.

Before initiation of CPB, the patients received intravenous heparin (300–500 units.kg\textsuperscript{-1} body weight) to achieve an ACT > 480 s. CPB was instituted via an ascending aortic cannula and a two-stage right atrial cannula. Before, during
(pump blood flow: 2.4L/min/m²), and after CPB, mean arterial pressure was adjusted to exceed 60mmHg. Cardiac arrest had been achieved with cold antegrade blood cardioplegia. Lactate-enriched Ringer’s solution was added to the CPB circuit to maintain reservoir volume when needed, and packed red blood cells would be added when hemoglobin concentration decreased to less than 7g.dl⁻¹.

After the patient rewarming to 37°C, separation from CPB, reversal of heparin by protamine sulfate (1:1), and sternal closure was achieved.

The anesthesiology intern prepared the study solution, bupivacaine 0.25% or normal saline, in the operating room. For carrying out the block bilaterally at the end of the surgery, the skin on either side of the sternum was prepared with povidone-iodine solution. Then, a linear ultrasound probe (Philips clear vue350, Philips Healthcare, Andover MAO1810, USA) was placed on the right and left sides at 3cm from the mid sternum. The subcutaneous tissue was identified; the intermediate plane: the pectoralis major muscle, the intercostal muscles, the ribs; the deep plane, the transverse thoracic muscle, the pleura, and the lung.

Block-group: had received bilateral TTMP block. After identifying the anatomical plane between the internal intercostal and the transversus thoracic muscles, a 22-gauge short bevel needle (Spinocan, B. Braun Melsungen AG, Germany) was inserted between the fourth and fifth ribs connecting at the sternum.

Correct needle placement was confirmed by visualizing the needle in the plane along its entire length and the tip of the needle between both muscles, then 1 mL of anesthetic liquid was introduced after negative aspiration. The TTMP block was completed by injection of 20 mL of 0.25% bupivacaine, and the same procedure was repeated on the other side. A local anesthetic indicated the appropriate injection spread deep to the costal cartilages and downward displacement of the pleura.

Non-block group: patients received sham block bilaterally with 20 mL of 0.9% saline had been injected on each side.

All patients were transported to ICU for postoperative management and care. Tracheal extubation was performed when the patient met the following criteria: awake/arousable, hemodynamically stable, no active bleeding, warm peripheries, and satisfactory arterial blood gas with an FiO₂ <0.5, pressure support on ventilator reduced to 10 Cm H₂O, Positive End Expiratory Pressure 5–7 CmH₂O, no electrolyte abnormalities, minimal inotropic support, or no escalation in inotropic support.

Postoperative analgesia in the ICU was carried out for both groups. All patients received IV fentanyl via patient-controlled analgesia (PCA) with (10µg.mL⁻¹, with a bolus of 15 µg, and lockout 10 minutes, maximum cumulative dose of 90µ.hr⁻¹ and no background dose). Before extubation, analgesia was given as nurse-controlled analgesia (NCA) with the same regimen, depending on the sudden rise in HR or MABP ≥20% of the baseline. Paracetamol 1gm was given every 8 hours for all patients. The total 24 h. opioid consumption was recorded.

Measured Parameters
The primary outcome was total fentanyl consumption (time frame: from ICU admission up to 24 hours). Secondary outcomes included: visual analog score (VAS) for sternal pain both during rest and with cough (ranging from 0 indicating no pain to 10 indicating extreme pain) measured at time intervals: 30 minutes, 1 hour, 3 hours, 6 hours, 12 hours, and 24h after extubation, time to extubation, the first analgesic request time, ICU length of stay and total length stay of the hospital.

Statistical Analysis
Before the study, the sample size was calculated. The primary outcome was the total fentanyl consumption during the first 24 hours after surgery. There were no previous studies when designing the study protocol. Therefore, we performed an external pilot study that included 9 patients in each group, with its results not included in the full-scale study. From this pilot, the total 24 hours postoperative fentanyl consumption (µg) was (371.7± 150.03 in the control group versus 260 ±110.45 in the block group with a mean reduction of 30%). The minimal sample size of patients was 31 in each group needed to get power level 0.90 and alpha level 0.05. The calculated sample size was increased by 10% to reach 35 in each group to allow for dropouts.

The collected data were organized, tabulated, and statistically analyzed using SPSS software statistical computer version 22 (SPSS Inc, USA). We used a two-sample t-test to compare the two groups’ mean values (age, BMI, fentanyl consumption)
and a Mann–Whitney U-test to compare medians for skewed endpoints (time of extubation, ICU stay, and hospital stay). The chi-squared test was used to determine significance, and qualitative data were presented as numbers and percentages (Sex, HTN, DM, and the operation type). Linear mixed models were used to account for repeated measures of pain scores. A fixed-effect model was used for the group and a random effect model for the subject. The first analgesic request time was assessed using the Kaplan–Meier estimator; Median time and (95% CI) were estimated, and the Log rank test was performed to compare study groups. A two-sided P-value of <0.05 was considered statistically significant.

**Results**

For this study, 76 patients were assessed for eligibility based on the inclusion and exclusion criteria. Six patients were excluded; two cases with EF below 35%, two patients declined to participate, and two due to stuck valves. The remaining 70 patients were randomly assigned into the block group, which received the block, and the non-block group, which control group (Figure 1).

There was no statistically significant difference between the two groups regarding the demographic characteristics and operative data (Table 1). In addition, there was no statistically significant difference between the two groups regarding the time elapsed in ICU or the hospital (Table 1).

Mean (SD) total fentanyl consumption in the first 24 hours was significantly lower in the block group than the no block group 205.7 (73.5) µg vs 390.9 (80.3) µg, with a mean difference of −185.143, 95% CI = −221.871 to −148.415; p < 0.0001 (Table 2).

The median estimate time to the first analgesic request was longer in the block group (14 hours, 95% CI = 12.17–15.84) than in the no block group (3 hours, 95% CI = 1.72–4.28), p < 0.0001 (Figure 2).

Using mixed effect model during the post-operative period (0.5–24 hours), at-rest pain scores were 1.86 units lower in the Block group than Non-block group (the estimate was −1.80, 95% CI = −2.14 to −1.45, t = −10.323 with p < 0.0001). Likewise, pain scores with cough were 3.29 units lower in the Block group than Non-block group (the estimate was −3.29, 95% CI = −3.80 to −2.77, t = −12.703, p < 0.0001) (Figure 3).

We did not report any block-related complications.

**Discussion**

Our study showed significantly decreased 24-hours fentanyl consumption with significantly prolonged time to first analgesic request in the block group than the non-block group. Also, we found reduced VAS scores of pain considerably both during rest and with cough after extubation in the block group than in the non-block group.

TTMP block can provide analgesia for post-sternotomy pain by targeting multiple anterior branches of the intercostal nerves (T 2–6). In their study, Aydin et al reported a significant decrease in 24-hour fentanyl consumption among adult patients who received TTMP block compared to the control group. Also, various studies have reported the efficacy of TTMP block in reducing opioid consumption after cardiac surgery among the pediatric population.9–11

On the other hand, Fujii et al reported that the 24-hour opioid requirement was similar in patients who received or did not receive the block, this can be attributed to; first: their study was a pilot study and included only 19 patients, which is a small number to declare a difference, second: lack of control of intraoperative and ICU opioid use which can affect postoperative pain scores and opioid requirement, furthermore, 60% of their patients underwent CABG with IMA harvesting. IMA harvesting will result in surgical disturbance of the TTMP block plane and uneven spread of the injectate between the desired thoracic levels. Patients may not benefit from the TTMP block on that side.13

We found that TTMP block significantly decreased VAS scores of pain at all-time intervals during the first 24 hours after extubation. Many studies have reported the same.9–11

However, Aydin et al reported lower VAS scores only in the first 12 hours after extubation with no difference between block and non-block groups at 24 hours; this can be attributed to the time of performing the block as they performed TTMP block before surgery.

We found no difference in extubation time between both groups. On the other hand, studies in pediatrics found a shorter time for extubation among patients who received the block.9–11 Exubation is not dependent only on pain control; other factors such as hemodynamic stability and complete reversibility of NMB can affect the extubation time.
We did not report any block-related complications. Also, in their studies, Aydin et al\(^8\) and Fujii et al\(^{12}\) wrote no block-related complications. However, in a case series of 299 consecutive TTMP block cases, Ueshima et al\(^{14}\) said two patients with infection around the injection site.

Our study found no significant differences between both groups in LOS. Also, Aydin et al\(^8\) and Cakmak and Isik\(^{11}\) have found the same. On the other hand, I.I. Abdelbaser and Mageed\(^7\) & Zhang et al\(^{10}\) showed a shorter LOS in the block groups compared with non-block groups.

Abbreviations: n, number; EF, ejection fraction

Figure 1 CONSORT flow diagram of the study population.
**Limitations**

Median sternotomy is not the only source of pain following cardiac surgery; chest tubes and visceral pain are considered other essential sources. In this study, we targeted only post-sternotomy pain; this can explain why patients in the block group requested opioids. Another limitation is the dependence on NCA based on clinical data for supplementary analgesia before extubation which could affect opioid consumption. Furthermore, all participants were Egyptians, and CABG surgeries were not included due to anatomical considerations, limiting our data’s generalizability. Again, the limited number of available clinical trials represented a difficulty for comparison. Despite these limitations, our results highlight the successful role of TTMP block in reducing pain and opioid consumption following median sternotomy.

**Table 1** Demographic Characteristics, Operative Data, and Length of Stay

<table>
<thead>
<tr>
<th></th>
<th>Group Block</th>
<th>Group No block</th>
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</thead>
<tbody>
<tr>
<td>Sample size, n</td>
<td>35</td>
<td>35</td>
</tr>
<tr>
<td>Mean age (SD) in (years)</td>
<td>38 (15)</td>
<td>39 (13)</td>
</tr>
<tr>
<td>Mean BMI (SD) in kg/m²</td>
<td>25 (4)</td>
<td>25 (3)</td>
</tr>
<tr>
<td>Sex, n (%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Male</td>
<td>22 (63)</td>
<td>22 (63)</td>
</tr>
<tr>
<td>Female</td>
<td>13 (37)</td>
<td>13 (37)</td>
</tr>
<tr>
<td>HTN, n (%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Yes</td>
<td>6 (17)</td>
<td>7 (20)</td>
</tr>
<tr>
<td>No</td>
<td>29 (83)</td>
<td>28 (80)</td>
</tr>
<tr>
<td>DM, n (%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Yes</td>
<td>4 (11)</td>
<td>6 (17)</td>
</tr>
<tr>
<td>No</td>
<td>31 (89)</td>
<td>29 (83)</td>
</tr>
<tr>
<td>The operation, n (%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>ASD</td>
<td>1 (3)</td>
<td>4 (11)</td>
</tr>
<tr>
<td>AVR</td>
<td>7 (20)</td>
<td>10 (29)</td>
</tr>
<tr>
<td>MVR</td>
<td>19 (54)</td>
<td>20 (57)</td>
</tr>
<tr>
<td>MVR &amp; AVR</td>
<td>6 (17)</td>
<td>1 (3)</td>
</tr>
<tr>
<td>MVR &amp; ASD</td>
<td>1 (3)</td>
<td>0 (0)</td>
</tr>
<tr>
<td>Triple valve</td>
<td>1 (3)</td>
<td>0 (0)</td>
</tr>
<tr>
<td>Median Time of extubation (IQR) in hours</td>
<td>5 (4–6)</td>
<td>6 (5–6)</td>
</tr>
<tr>
<td>Median ICU stay(IQR) in hours</td>
<td>36 (36–48)</td>
<td>36 (36–60)</td>
</tr>
<tr>
<td>Median Hospital stay(IQR) in days</td>
<td>4 (3–5)</td>
<td>4 (3–4)</td>
</tr>
</tbody>
</table>

**Table 2** Total Fentanyl Consumption in the First 24 Hours

<table>
<thead>
<tr>
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<th>Block Group</th>
<th>Non-Block Group</th>
<th>Mean Difference 95% CI</th>
<th>P-value</th>
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<td>Mean total fentanyl consumption (SD) in (µg)</td>
<td>205.7 (73.5)</td>
<td>390.9 (80.3)</td>
<td>−185.143 (−221.871 to −148.415)</td>
<td>&lt;0.0001*</td>
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</tbody>
</table>

**Abbreviations:** SD, standard deviation; n, number; ASD, atrial septal defect; AVR, aortic valve replacement; BMI, body mass index; DM, diabetes mellitus; HTN, hypertension; ICU, intensive care unit; MVR, mitral valve replacement; IQR, interquartile range.

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**Abbreviations:** SD, standard deviation; n, number; SD, *statistically significant.

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**Conclusion**

TTMP block successfully reduces postoperative opioid consumption, prolongs time to first analgesic request, and decreases pain scores. It has an opioid-sparing effect and can be used as a part of a multimodal analgesia regimen in a patient undergoing open-heart surgeries via median sternotomy. We recommend further clinical trials over a large-scale population to ensure the clinical role of TTMP block as apart from the multimodal analgesic regimen. Also, further studies are still requested to evaluate the possibility and efficacy of catheter placement for multiple injections.
Figure 2 The time to the first analgesic request.

Figure 3 The VAS score at rest and with cough.

Abbreviations: VAS, visual analog score; h, hours.
Registration
This study is registered on ClinicalTrials.gov (NCT04116554; principal investigator: Mohamed Ahmed Hamed; date of registration: October 4, 2019).

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Disclosure
The authors declare no conflicts of interest in this work.

References