INTRODUCTION

Indications for the strategy known as the open abdomen technique (OAT) include the treatment of severe abdominal sepsis, the prevention or treatment of intra-abdominal hypertension, and the need for damage control in cases of intra-abdominal bleeding (1). However, the treatment of diffuse peritonitis, despite advances, remains challenging (2,3).

It is known that temporary closure of the abdomen with the use of traditional dressings is associated with increased mortality and morbidity compared with closure of the abdomen and relaparotomy on demand (4-7). However, a recent study suggested that the OAT with closure of the abdomen using dressings that provide negative pressure is associated with favorable results (1).

With the advent of vacuum therapy, the use of laparostomy to treat diffuse peritonitis has gained traction (1,8). Despite these advances, patients treated with OAT are submitted to many procedures, and extensive time is generally spent in the intensive care unit because, regardless of the technique employed, the laparostomy device must be changed several times until the patient recovers.

Another modality of treatment of diffuse peritonitis is continuous peritoneal lavage (9), which aims to decrease the number of procedures and maintain continuous washing concurrently with intensive care support. However, the concept of continuous peritoneal lavage is technically difficult and has been associated with controversial results (3).

No technique is currently available that reduces the number of procedures and the stress of dressing removal in
OAT. Additionally, no technique combines the benefits of vacuum laparostomy with continuous peritoneal lavage. The concept of irrigation and aspiration through the vacuum system is currently used only for superficial wounds (10,11).

Thus, in this experiment, we evaluated the performance of a model of continuous peritoneal lavage taking advantage of the features and benefits of vacuum laparostomy, and we analyzed the viability of continuously maintaining fluid infusion in the peritoneum through an intermittent vacuum laparostomy dressing.

■ MATERIAL AND METHODS

Study design and ethical statement

The research protocol was approved by the Research Ethics Committee for the use of animal models at our institution (Protocol No. 086/12). All animals were handled according to the principles of the National Institutes of Health (1985) and The American Physiological Society (1995) for the care, handling and use of laboratory animals. An experimental study was carried out in which four Landrace and Large White swine weighing between 20 and 30 kg underwent the experimental procedure. Exclusion criteria or animals were a plasma hemoglobin concentration lower than 9 mg/dL, abnormal blood gas values at baseline, clinical signs of infection, and early hemodynamic deterioration (blood loss <500 mL associated with mean arterial pressure (MAP) <60 mmHg in the first 10 minutes. After the experimental procedure, the animals were sacrificed with an overdose injection of sodium phenobarbital.

Anesthetic protocol, instrumentation and monitoring

The preparation of the animals included fasting for 12 hours with access to water ad libitum. Then, the animals were sedated with an intramuscular injection of ketamine (5 mg kg⁻¹) and midazolam (0.25 mg kg⁻¹), and anesthesia was induced with i.v. administration of propofol (5 mg kg⁻¹).

After endotracheal intubation, anesthesia was maintained with isoflurane (1.5% by volume), and pancuronium bromide (5 mg kg⁻¹ min⁻¹) was used to ensure muscle paralysis, which was monitored throughout the experiment.

The ventilation was adjusted for volume control of the following parameters: FiO₂ of 40%, 8 mL kg⁻¹, 5 cmH₂O of positive end-expiratory pressure (PEEP) and a respiratory rate adequate to maintain PaCO₂ between 30 mmHg and 35 mmHg. A heating blanket was used to maintain the temperature of the animals at approximately 38°C.

A multiparametric monitor (IntelliVue MP40; Phillips, Boeblingen, Germany) was used to evaluate heart rate (HR) and rhythm throughout the procedure.

The right internal jugular vein was punctured for the introduction of a continuous pulmonary artery catheter coupled with a mixed oxygen venous saturation monitor (SvO₂) connected to a cardiac output monitor. A catheter was introduced via the pressure transducer connection and was guided by analyzing the pressure curves of AD, VD, AP and POAP. The midpoint between the anterior and posterior thoracic walls was taken as the zero reference point for pressure measurements. The intravascular catheters were adjusted to the atmospheric parameters.

A standard Foley catheter was positioned in the bladder and connected to a pressure transducer for assessment of intra-abdominal pressure (IAP).

Technique

The animals were submitted to a continuous peritoneal lavage technique associated with vacuum peritoneostomy, as previously described (12). They were submitted to laparostomy through which a 4.8 mm multiperforated tube was placed along each flank and exteriorized in the lower left and lower right quadrants. An adapted vacuum dressing was placed in the laparostomy (VAC® Dressing, KCI), and an intermittent negative pressure of 125 mmHg was maintained with vacuuming every 30 minutes for 30 minutes.

Hemodynamic and biochemical parameters and bladder pressure values were collected at baseline (BL), 6 hours (T6), 12 hours (T12), 18 hours (T18), 24 hours (T24) and 36 hours (T36).

CT image evaluation

The animals received 60 ml/kg PDS through the two multiperforated tubes and were submitted to CT without the use of intravenous contrast.

To quantify the fluid present in the cavity, a scoring system called the abdominal fluid distribution score (AFDS) was used. The AFDS assesses the presence or absence of fluid and the amount of fluid visualized on CT images. The abdominal cavity of the animal was divided into 6 regions: the right costophrenic recess (RCFR), the left costophrenic recess (LCFR), the subhepatic region (SH), between loops (BL), the right iliac fossa (RIF) and the left iliac fossa (LIF).

For AFDS analysis, one point was assigned for liquid presence, and two points were assigned if the presence of liquid was greater than 1 cm in the image for each region. The minimum score was zero, i.e., no fluid in any space, and the maximum score was twelve corresponding to the presence of fluid in all spaces with a thickness greater than 1 cm in the CT image. A score of 6 was considered sufficient to perform washing in this model.

System stability and physiological effects of washing with PDS

Peritoneostomy stability with intermittent infusion of fluids, system resistance to obstructions and leakage, water balance, hemodynamic parameters (mean arterial pressure, HR, IAP, diuresis, CO) and biochemical parameters (pH, lactate, sodium, potassium, hematocrit and SatO₂) were analyzed every 6 hours.

Statistical analysis

Data were analyzed using the SigmaStat program for Windows and are presented as medians. The Wilcoxon paired test was used to compare hemodynamic and biochemical variables collected at the beginning and end of the experiment.

■ RESULTS

Disposition of the peritoneal lavage fluid in the abdominal cavity

Through analysis of CT images, it was possible to verify efficiency of the system in occupying all spaces. Table 1 shows the scoring results using the AFDS. The results were considered adequate according to established criteria.

Although previously positioned in the right and left flanks, the multiperforated tubes left their original position in all animals. However, this movement of the tubes did not
affect the distribution of fluids within the abdominal cavity (Figures 1(A-D)).

Stability and physiological effects

Stability of the peritoneostomy was maintained during the procedure, with no leakage, catheter obstruction or changes in IAP. The water balance evaluation showed that the volume aspirated by peritoneostomy at the end of the experiment was similar to the volume infused (10165 ± 235.73 mL versus 9073.5 ± 1496.35 mL, p=0.25). There were no significant changes in hemodynamic or biochemical parameters (Tables 2 and 3).

## DISCUSSION

The continuous peritoneal lavage strategy as a treatment for abdominal sepsis is supported by the idea that continuous irrigation of the abdominal cavity washes out inflammatory mediators and contaminants, thus resolving the infection. This theory has been studied since the 1960s, although there is little evidence supporting this approach in clinical and scientific literature (13,14). It has also been postulated that current techniques and more aggressive washing may cause more harm than good (9).

Clinical and experimental controversial and unfavorable results of continuous peritoneal lavage may be associated with the approaches that are currently used. These approaches are not capable of washing the entire cavity and/or do not effectively clear contaminants and inflammatory mediators, since they do not completely irrigate the abdominal cavity. Intracavitary catheters are generally positioned, and it is expected that irrigation, without proper removal of the infused fluid, will resolve the intra-abdominal infection. In addition, the abdominal cavity is kept closed, which contributes to increased abdominal pressure. A few studies have evaluated peritoneostomy with peritoneal lavage, but there are no comparative studies (15,16).

Finally, the technical difficulties of continuous peritoneal lavage are associated with maintaining the integrity of the system. Most methods use the same fluid infusion route to aspirate the lavage, causing leaks, obstructions, contamination and infection of the remaining volume (3).

In a recent publication (12), we described a continuous peritoneal lavage model associated with vacuum peritoneostomy that aimed to solve the problems observed in previous peritoneal lavage techniques. The model was able

Table 1 - Abdominal Fluid Distribution Score assessed from CT images.

| Animals | RCFR | LCFR | SH | BL | RIF | LIF | Total |
|---------|------|------|----|----|-----|-----|-------|
| 1       | 1    | 2    | 2  | 1  | 2   | 2   | 10    |
| 2       | 1    | 1    | 2  | 1  | 2   | 2   | 9     |
| 3       | 1    | 1    | 2  | 2  | 2   | 2   | 10    |
| 4       | 1    | 1    | 2  | 2  | 2   | 2   | 9     |

RCFR: Right costophrenic recess; LCFR: Left costophrenic recess; SH: Subhepatic; BL: Between loops; RIF: Right iliac fossa; LIF: Left iliac fossa.

Figure 1 - (A) Pig 1, (B) Pig 2, (C) Pig 3 and (D) Pig 4. The red arrows indicate the presence of liquid in all spaces evaluated.
Continuous peritoneal lavage

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Table 2 - Hemodynamic variables of animals submitted to peritoneal lavage with PDS.

| Variable    | BL (median) | T6 (median) | T12 (median) | T18 (median) | T24 (median) | T36 (median) | p-value* |
|-------------|-------------|-------------|-------------|-------------|-------------|-------------|---------|
| MAP (mmHg)  | 66.5        | 64.5        | 62.5        | 61.5        | 68.5        | 65          | 0.42    |
| HR (bpm)    | 91.5        | 101         | 87.5        | 106         | 103         | 97.5        | 0.71    |
| IAP (mmHg)  | 5.5         | 5.5         | 8           | 8           | 17.5        | 13          | 0.07    |
| Diuresis (mL)| 350         | 280         | 442         | 566         | 642         | 900         | 0.07    |
| CO (L/min)  | 2.7         | 3           | 2.65        | 2.8         | 2.45        | 2.95        | 0.27    |

Table 3 - Biochemical variables of animals submitted to peritoneal lavage with PDS.

| Variable    | BL (median) | T6 (median) | T12 (median) | T18 (median) | T24 (median) | T36 (median) | p-value* |
|-------------|-------------|-------------|-------------|-------------|-------------|-------------|---------|
| pH          | 7.44        | 7.49        | 7.49        | 7.49        | 7.48        | 7.47        | 0.14    |
| Lactate (nmol/L) | 1.15      | 0.82        | 1.07        | 1.02        | 0.77        | 0.74        | 0.14    |
| Na (nmol/L)  | 138.50      | 133.5       | 136         | 134         | 133.50      | 134.50      | 0.07    |
| K (nmol/L)   | 3.77        | 4.56        | 4.25        | 4.21        | 4.05        | 4.31        | 0.07    |
| Ht (%)       | 27.40       | 30.60       | 29.95       | 29.70       | 33.85       | 29.90       | 0.59    |
| SatO2 (%)    | 99.50       | 99.80       | 99.40       | 99.50       | 99.50       | 97.95       | 0.99    |

Table 4 - Results of hemodynamic and biochemical variables of animals submitted to peritoneal lavage with PDS.

PDS: Peritoneal dialysis solution; MAP: Mean blood pressure; HR: Heart rate; IAP: intra-abdominal pressure; CO: Cardiac output; * Paired Wilcoxon test (T0XT36).

Continuous peritoneal lavage with PDS associated with vacuum peritoneostomy may be useful in the treatment of diffuse peritonitis.

**CONCLUSION**

Continuous peritoneal lavage with PDS associated with vacuum peritoneostomy is technically feasible and maintains physiological parameters within the normal range. Thus, clinical studies should be conducted to evaluate the use of this technique in the treatment of diffuse peritonitis.

**AUTHOR CONTRIBUTIONS**

Rodrigues AC and Utiyama EM were responsible for the conception, design, intellectual and scientific content of the study. Santos LC and Otsuki DA were responsible for animal procedures, management, monitoring and data collection. Saad KR and Saad PF were responsible for statistics and manuscript writing. Montero EFS, Rasslan S, Utiyama EM were responsible for interpretation of the data, discussion and ensuring adequacy of the model for clinical applications. All authors revised the final text of the manuscript and contributed to the final version.

**REFERENCES**

1. Demetriades D. Total management of the open abdomen. Int J. 2013;9(Suppl.1):17-24, https://doi.org/10.1111/1472-4481.X.2012.01018.x.
2. Buijk SE, Bruining HA. Future directions in the management of tertiary peritonitis. Intensive Care Med. 2002;28(8):1024-9, https://doi.org/10.1007/s00134-002-1363-6.
3. Van Goor H. Interventional management of abdominal sepsis: when and how. Langenbecks Arch Surg. 2002;387(5-6):191-200, https://doi.org/10.1007/s00423-002-0309-7.
4. Leppäniemi AK. Laparostomy: why and when? Crit Care. 2010;14(2):216, https://doi.org/10.1016/j.ccrs.2010.10.001.
5. Robledo FA, Luque-de-León E, Suárez R, Sánchez P, de-la-Fuente M, Vargas A, et al. Open versus closed management of the abdomen in the surgical treatment of severe secondary peritonitis: a randomized clinical trial. Surg Infect. 2007;8(3):63-72, https://doi.org/10.1089/sur.2006.8.016.
6. Christou NV, Barie PS, Dellinger EP, Waymack JP, Stone HH. Surgical Infection Society intra-abdominal infection study. Prospective evaluation of management techniques and outcome. Arch Surg. 1993;128(2):198-9, https://doi.org/10.1001/archsurg.1993.00134050070011.
7. Adjíns AL, Robbins J, Villaíla M, Bendick P, Stanley CJ. Open abdomen management of intra-abdominal sepsis. Am Surg. 2004;70(2):137-40.
8. Perathoner A, Klaus A, Mühlmann G, Oberwalder M, Margreiter R, Kafka-Ritsch R. Damage control with abdominal vacuum therapy (VAC) to manage perforated diverticulitis with advanced generalized peritonitis.
peritonitis - a proof of concept. Int J Colorectal Dis. 2010;25(6):767-74, https://doi.org/10.1007/s00384-010-0887-8.

9. Platell C, Papadimitriou JM, Hall JC. The influence of lavage on peritonitis. J Am Coll Surg. 2000;191(6):672-80, https://doi.org/10.1016/S1072-7515(00)00726-2.

10. Ugaki S, Kasahara S, Arai S, Takagaki M, Sano S. Combination of continuous irrigation and vacuum-assisted closure is effective for mediastinitis after cardiac surgery in small children. Interact Cardiovasc Thorac Surg. 2010;11(3):247-51, https://doi.org/10.1510/icvts.2010.235903.

11. D’Hondt M, D’Haeninck A, Dedrye L, Penninckx F, Aerts R. Can vacuum-assisted closure and instillation therapy (VAC-Instill™ therapy) play a role in the treatment of the infected open abdomen? Tech Coloproctol. 2011; 15(1):75-7, https://doi.org/10.1007/s10151-010-0662-4.

12. Rodrigues AC, Santos LCD, Otsuki DA, Saad KR, Saad PF, Montero EFS, et al. Animal model of continuous peritoneal lavage with vacuum peritoneostomy. Acta Cir Bras. 2017;32(6):467-74, https://doi.org/10.1590/s0102-8630201700000007.

13. Shams WE, Hanley GA, Orvik A, Lewis N, Shurbaji MS. Peritoneal lavage using chlorhexidine gluconate at the end of colon surgery reduces postoperative intra-abdominal infection in mice. J Surg Res. 2015;195(1):121-7, https://doi.org/10.1016/j.jss.2015.01.030.

14. Glover JL, Atkins P, Lempke RE. Evaluation of peritoneal lavage therapy for peritonitis. J Surg Res.1997;9:3:134-5, https://doi.org/10.1016/S0022-4804(99)00130-9.

15. Losasso J, Kjosev K. Palisade dorsoventral lavage for neglected peritonitis. Am J Surg. 1997;173(2):134-5, https://doi.org/10.1016/S0002-9610(96)00411-4.

16. Dittrich K, Kriwanek S, Armbruster C. The Incise Pouch: a helpful device in continuous postoperative dorso-ventral lavage procedures. Langenbecks Arch Chir. 1993;378:185-7, https://doi.org/10.1007/BF00184470.

17. Breborowicz A, Oreopoulos DG. Is normal saline harmful to the peritoneum? Perit Dial Int. 2005;25 Suppl 4:S67-70.

18. Pohubińska A, Winckiewicz M, Staniszewski R, Breborowicz A, Oreopoulos DG. Time to reconsider saline as the ideal rinsing solution during abdominal surgery. Am J Surg. 2006;192(3):281-5, https://doi.org/10.1016/j.amjsurg.2005.05.047.

19. Carr JA. Abdominal compartment syndrome: a decade of progress. J Am Coll Surg. 2013;216(1):135-46, https://doi.org/10.1016/j.jamcollsurg.2012.09.004.