

Effect of hyperosmolar sodium lactate infusion on haemodynamic status and fluid balance compared with hydroxyethyl starch 6% during the cardiac surgery

Address for correspondence:
Dr. Cindy Elfira Boom,
Department of Anesthesiology,
Harapan Kita National
Cardiovascular Center, Jl.
Letjend. S. Parman Kav. 87,
Jakarta 11420, Indonesia.
E-mail: cindynugroho@yahoo.
com

**Cindy Elfira Boom, Poernomo Herdono, Chairil Gani Koto, Sjamsul Hadi,
I Made Adi Permana**

Department of Anaesthesia and Intensive Care, National Cardiovascular Center, Harapan Kita Hospital,
Jakarta, Indonesia

ABSTRACT

Background and Aim: No solution has been determined ideal for fluid therapy during cardiac surgery. Previous studies have shown that hyperosmolar sodium lactate (HSL) infusion has improved cardiac performance with smaller volume infusion, which resulted in negative fluid balance. This study compared the effects between a patent-protected HSL infusion and hydroxyethyl starch (HES) 6% on haemodynamic status of the patients undergoing cardiac surgery. **Methods:** In this open-label prospective controlled randomized study, patients were randomly assigned to receive loading dose of either HSL or HES 6%, at 3 mL/kgBW within 15 min, at the beginning of surgery. Haemodynamic parameters and fluid balance were evaluated, while biochemical parameters and any adverse effect were also recorded. Haemodynamic and laboratory parameters were analyzed through repeated measures analysis of variance. Statistical assessment of fluid management was carried out through Student *t*-test. All statistical analyses were performed using the statistical package for the social sciences® version 15, 2006 (SPSS Inc., Chicago, IL). **Results:** Out of 100 enrolled patients in this study (50 patients in each arm), 98 patients were included in analysis (50 in HSL group; 48 in HES group). Cardiac index increased higher in HSL group ($P = 0.01$), whereas systemic vascular resistance index decreased more in HSL than HES group ($P = 0.002$). Other haemodynamic parameters were comparable between HSL and HES group. Fluid balance was negative in HSL group, but it was positive in HES group (-445.94 ± 815.30 mL vs. $+108.479 \pm 1219.91$ mL, $P < 0.009$). **Conclusion:** Administration of HSL solution during the cardiac surgery improved cardiac performance and haemodynamic status better than HES did.

Key words: Cardiac surgery, fluid resuscitation, hydroxyethyl starch, hyperosmolar sodium lactate

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INTRODUCTION

Maximising the cardiac output (CO) by fluid infusion benefits patients undergoing cardiac surgery,^[1-3] but they may not tolerate large volume of fluid due to impaired cardiac performance. Hence, fluid resuscitation without or with minimal risks of fluid excess might be beneficial.^[4,5] Colloids are widely used in cardiac perioperative setting,^[4,6,7] however,

higher cost and potential risks of colloids are still unresolved. Hypertonic solution infusion has been shown to benefit cardiac surgery patients^[4,8-12] and was associated with a higher excretion of body fluid excess when compared with colloid infusion.^[13]

Perioperative administration of lactate based hyperosmolar solutions benefited cardiac surgery patients by improving cardiac performance, oxygen

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delivery and inducing negative fluid balance.^[14,15] Lactate was suspected to contribute to these benefits as per the findings of previous studies.^[16-21] We conducted this study to evaluate the effect of hyperosmolar sodium lactate (HSL) infusion on haemodynamics and fluid balance in comparison with 6% hydroxyethyl starch (HES) infusion.

METHODS

This prospective, randomised, open-labelled study aimed to evaluate the efficacy and the safety of a scientifically-formulated and patent-protected HSL solution (Totilac®), manufactured by Finusolprima Farma Indonesia (for Innogene Kalbiotech Pte Ltd) compared to HES 6% (Voluven™, Fresenius Kabi) during coronary artery bypass grafting (CABG) surgery. The composition of each solution is described in Table 1. This study was approved by institutional ethical committee.

Male and female patients, aged 18-75 years undergoing CABG surgery were enrolled. We excluded the patients who needed combined operations and intra-aortic balloon pump, patients with severe arrhythmia (ventricular tachycardia, atrial flutter with rapid response, heart block), the presence of severe haemodynamic imbalance, severe bleeding and/or re-operation, liver dysfunction as indicated by serum glutamic oxaloacetic transaminase and serum glutamic pyruvic transaminase >2 times normal value) and renal failure (creatinine level >180 µmol/L). Signed informed consent was obtained from all eligible patients or the next of kin if the patient was not able to comprehend or unable to accept and sign informed consent.

Patients underwent routine monitoring including 5-lead electrocardiography, radial, central venous pressure (CVP), pulmonary artery (PA) catheters,

pulse oximetry, blood, urine and temperature monitoring. Following pre-medication with intravenous midazolam (2.5-5 mg), anaesthesia was induced with 3-5 mg/kgBW propofol, titrated and adjusted based on patients haemodynamic status and clinical condition and 1-2 µg/kgBW sufentanil. A dose of 0.1-0.2 mg/kgBW pancuronium bromide was administered for intubation.

Anaesthesia and haemodynamic stability during surgery was maintained with continuous infusion of 1.0-1.5 mg/kgBW of propofol as baseline hypnosis, coupled with titration of inhalational anaesthesia agent (sevoflurane). Then propofol was continued as post-operative sedation with patient warming and improvement in CO. Intravenous bolus of sufentanil was administered in case of poor analgesia and/or increase in blood pressure or heart rate at 30% or more. Nitroglycerin was continuously infused only for a tight indication, such as for patients with left main coronary artery occlusion or low ejection fraction (<40%) with multiple coronary occlusions, or if there were any signs of revascularisation injury. The infusion rate was modulated according to the blood pressure at 0.5 to 2 µg/kgBW/h. CO/cardiac index (CI), pulmonary vascular resistance (PVR)/pulmonary vascular resistance index (PVRI) and systemic vascular resistance (SVR)/systemic vascular resistance index (SVRI) were also measured and monitored intermittently by PA catheter to ensure that the haemodynamic status was normal during the infusion.

For cardiopulmonary bypass (CPB), the extracorporeal circuit consisted of a roller pump (Sarns 8000®) and a membrane oxygenator (Capiiox Sx®). The pump flow was 2.4 L/min/m². During CPB, patients were cooled to 31-32°C and received intermittent antegrade normothermic blood cardioplegia, a mixture of 400-600 mL of oxygenated blood with graduated doses of potassium-magnesium solution. Standard systemic heparinisation (3 mg/kgBW) was performed and an activated clotting time of greater than 480 s was maintained during CPB. Heparin was neutralized with protamine sulphate on discontinuation.

Patients were randomised by block permutation with block size of 3. Randomisation was conducted by an independent nurse and random allocation was concealed within a thick envelope. Study drug was administered by a nurse according to random allocation; hence the investigator was blind about the treatment. Similar dose of HSL or HES solutions

Table 1: Composition and profile of the HES 6% solution and HSL solution

Solute contents	HSL		HES 6%
	mmol/L	g/L	g/L
Na ⁺	504.15	11.5	-
K ⁺	4.02	0.16	-
Ca ²⁺	1.36	0.050	-
Cl ⁻	6.74	0.24	-
Lactate ⁻	504.15	44.92	-
Hydroxyethyl starch (HES 130/0.4)	-	-	60
NaCl	-	-	9
Calculated total osmolarity (mosm/L)	1020.42		N/A

HSL – Hyperosmolar sodium lactate; HES – Hydroxyethyl starch

were administered at 3 mL/kgBW within 15 min at the beginning of surgery (after induction and fasting fluid replacement). Additional fluid were given as needed, which was managed similarly in both groups. Types and amount of additional fluids administered were recorded.

Patients were intensively monitored during surgical procedure. The haemodynamic parameters were recorded on baseline and immediately after study fluid administration including heart rate (HR), systolic and diastolic blood pressure, mean arterial pressure (MAP), CO, CI, CVP, pulmonary capillary wedge pressure (PCWP), mean pulmonary arterial pressure, SVR and PVR. SVRI and PVRI were subsequently calculated using standard formulae. Laboratory parameters including arterial pH, PaO₂, PaCO₂, bicarbonate (HCO₃), haemoglobin and haematocrit levels, blood electrolytes (sodium, potassium, chloride, magnesium), blood glucose and blood lactate were recorded. Intra-operative fluid input, urine output, bleeding volume and fluid balance were also measured at the end of surgery. Any adverse event during the study period was documented.

Sample size calculation in this study was based on the anticipated minimal difference of CI between control and treatment group (delta value). Previous trials in cardiac surgery showed that hypertonic solution would result CI differences of 0.3 L/min/m² compared to isotonic crystalloid solution.^[15] Based on a two-tailed *t*-test with 5% of significance level and assuming standard deviation of 0.4, with 10% drop-out and power 80%, we required a total of 100 patients (50 in HSL group and 50 in HES group).

Statistical assessment of haemodynamic status and laboratory parameters was carried out through repeated measures analysis of variance (ANOVA). The within-group (HSL vs. HES) changes between baseline and loading dose was evaluated using ANOVA. The multivariate analysis was conducted to see the influence of each covariate on CI. Statistical assessment of fluid management was carried out through Student *t*-test. All statistical analyses were performed using statistical package for the social sciences® version 15, 2006 (SPSS Inc., Chicago, IL).

RESULTS

One hundred patients undergoing cardiopulmonary bypass graft (CABG) surgery, 50 in HSL group and 50 in

HES group, were enrolled in this study, however, only 98 (50 patients in HSL group and 48 patients in HES group) were included for analysis. Two patients in HES group were not included in the analysis due to incomplete data (1 patient) and protocol violation (1 patient). The demographics and baseline between HSL and HES group were comparable [Table 2].

Systolic and diastolic blood pressure, MAP, CVP, PCWP, PVR/PVRI showed similar changes between the two groups two groups (*P* > 0.05), while the increase of heart rate (*P* = 0.024), CO (*P* = 0.002), CI (*P* = 0.001) were higher in HSL than HES group [Table 3]. SVR/SVRI were significantly lower in HSL than HES group (*P* = 0.03 and *P* = 0.002 respectively) [Table 3]. The analysis evaluating the influence of co-variables

Table 2: Baseline characteristics and intra-operative data

Characteristics	HSL n=50	HES n=48	<i>P</i>
Age (years)	56.49±8.42	56.00±6.57	0.747
Height (cm)	164.51±7.44	164.73±6.19	0.873
Weight (kg)	68.12±10.89	67.93±9.74	0.929
Body mass index (kg/m ²)	25.14±3.50	24.99±3.37	0.835
Ejection fraction (%)	53.58±14.15	53.81±15.35	0.938
Plasma urea (mg/dL)	31.96±13.82	28.72±11.14	0.198
Creatinine (mg/dL)	1.10±0.44	1.05±0.31	0.521
SGOT (IU/L)	19.47±7.45	21.81±6.11	0.102
SGPT (IU/L)	26.69±14.89	31.62±11.47	0.078
Systolic blood pressure (mmHg)	112.74±13.50	113.94±19.20	0.722
Diastolic blood pressure (mmHg)	60.04±8.49	60.17±9.63	0.945
Mean arterial blood pressure (mmHg)	76.49±10.25	76.51±12.93	0.993
Heart rate (beats/min)	63.02±12.26	63.13±12.73	0.965
Cardiac output (L/min)	3.70±0.94	3.82±1.16	0.571
Cardiac index (L/min/m)	2.12±0.50	2.18±0.58	0.583
PAM (mmHg)	16.74±4.99	17.15±6.46	0.728
PCWP (mmHg)	10.21±7.79	11.09±5.71	0.517
CVP (mmHg)	7.30±3.22	7.19±2.94	0.86
SVR (dyne×s/cm ⁵)	1,567.57±453.57	1,495.46±435.74	0.422
SVRI (dyne×s/cm ⁵)	2641.26±648.321	2528.21±673.89	0.399
PVR (dyne×s/cm ⁵)	180.64±115.60	141.75±97.58	0.071
PVRI (dyne×s/cm ⁵)	293.48±136.06	244.74±170.36	0.123
CPB time (min)	101.45±32.29	115.73±32.89	0.052
AOX time (min)	73.90±24.48	78.02±27.42	0.478
Operation duration (min)	372.47±122.82	380.61±82.08	0.733

BMI – Body mass index; EF – Ejection fraction; SGPT – Serum glutamic pyruvic transaminase; SGOT – Serum glutamic-oxaloacetic transaminase; PAM – Mean pulmonary artery pressure; PCWP – Pulmonary capillary wedge pressure; CVP – Central venous pressure; SVR – Systemic vascular resistance; SVRI – Systemic vascular resistance index; PVR – Pulmonary vascular resistance; PVRI – Pulmonary vascular resistance index; CPB – Cardio pulmonary bypass; AOX – Aortic cross clamping; HSL – Hyperosmolar sodium lactate; HES – Hydroxyethyl starch; SD – Standard deviation. Values are mean±SD

Table 3: Haemodynamic evolutions

Haemodynamic parameters	Study drugs	Baseline	Loading 1	Drug and dose interaction (<i>P</i> value)
SBP (mmHg)	HSL	111.78±12.148	114.66±13.819	0.588
	HES	112.87±17.358	117.67±17.261	
DBP (mmHg)	HSL	60.14±8.512	60.92±10.515	0.092
	HES	59.77±9.943	64.27±11.797	
MAP (mmHg)	HSL	76.44±10.015	80.00±13.251	0.591
	HES	75.52±12.211	80.58±13.162	
HR (beats/min)	HSL	62.94±11.898	69.80±16.494	0.024
	HES	63.73±12.927	65.96±14.518	
CO (L/min)	HSL	3.62±1.062	4.46±1.606	0.002
	HES	3.80±1.163	4.05±1.039	
CI (L/min/m ²)	HSL	2.10±0.540	2.60±0.862	0.001
	HES	2.18±0.588	2.34±0.534	
PAM (mmHg)	HSL	17.88±9.286	18.20±4.986	0.480
	HES	16.75±5.529	18.13±6.661	
PCWP (mmHg)	HSL	10.74±7.534	10.62±4.304	0.260
	HES	10.83±4.974	11.85±6.325	
CVP (mmHg)	HSL	7.70±3.059	8.24±2.745	0.473
	HES	7.02±2.556	7.96±3.235	
SVR (dyne.s/cm ⁵)	HSL	1624.38±533.426	1400.44±488.419	0.003
	HES	1481.04±403.651	1492.04±391.092	
SVRI (dyne.s/cm ⁵)	HSL	2707.66±741.399	2344.26±727.985	0.002
	HES	2442.75±733.503	2453.06±643.469	
PVR (dyne.s/cm ⁵)	HSL	170.50±117.586	148.56±119.718	0.224
	HES	131.17±94.226	136.62±85.376	
PVRI (dyne.s/cm ⁵)	HSL	291.34±193.020	252.02±187.005	0.366
	HES	233.15±156.058	226.58±155.062	

SBP – Systolic blood pressure; DBP – Diastolic blood pressure; MAP – Mean arterial pressure; HR – Heart rate; CO – Cardiac output; CI – Cardiac index; PAM – Pulmonary artery pressure; PCWP – Pulmonary capillary wedge pressure; CVP – Central venous pressure; SVR – Systemic vascular resistance; SVRI – Systemic vascular resistance index; PVR – Pulmonary vascular resistance; PVRI – Pulmonary vascular resistance index; SD – Standard deviation; HSL – Hyperosmolar sodium lactate; HES – Hydroxyethyl starch; ANOVA – Analysis of variance. Results are expressed as means±SD. By repeated measure ANOVA; statistically significant for *P*<0.05*

such as systolic and diastolic pressure, CVP, SVRI, PVRI, MAP, pH, HCO₃ and lactate on CI changes due to the treatment showed that SVRI was the most significant parameter related to the HSL treatment and also to the observed changes in CI between two treatment groups (*P* < 0.001). PVRI and HCO₃ levels level also influenced the changes of CI between two groups (*P* = 0.029 and 0.031, respectively).

The total fluid intake was not different between HSL and HES group although the amount of crystalloid infusion was lower in HSL than HES group (0.024) [Table 4]. The cumulative urine output was higher in HSL than HES group although it was not significant. The amount of urine output and blood loss was comparable between both groups. Hence, the total fluid output throughout the surgery was comparable between HSL and HES group. The fluid balance was more negative in HSL group (-445.94 ± 815.30), whereas the fluid balance was still positive in HES group (108.479 ± 1219.91); the difference was significant between the two groups (*P* = 0.009).

Baseline data of biochemical parameters was similar between HSL and HES group (*P* > 0.05). The changes of serum potassium, magnesium, haemoglobin, PaO₂, PaCO₂, SaO₂ levels, glucose level before and after study drug infusion were comparable between HSL and HES group (*P* > 0.05). Serum sodium level was higher in HSL than HES group (*P* < 0.0001). Blood lactate and pH were increased higher in HSL than HES group (*P* < 0.001). Arterial HCO₃ level increased in HSL group but decreased in HES group, the difference between both groups was significant (*P* < 0.001). Base excess reached normal level in HSL group, whereas in HES group it decreased to a more negative base excess, the difference between both groups was obviously very significant (*P* < 0.001) [Table 5].

The number of patients requiring vasodilators and or inotropes was numerically less in HSL than HES group, although it was not statistically different [Table 6].

No adverse events related to study drug infusion was noticed in both groups during the study period.

Table 4: Laboratory parameters before and after study drugs administration

Laboratory parameters	Study drugs	Baseline	Loading 1	Drug and dose interaction (P value)
Na ⁺	HSL	138.80±2.799	141.04±2.672	<0.0001
	HES	139.229±2.904	139.02±3.056	
K ⁺	HSL	3.65±0.333	3.66±0.377	0.687
	HES	3.64±0.379	3.63±0.365	
Ca ⁺⁺	HSL	1.11±0.207	1.03±0.199	0.085
	HES	1.15±0.199	1.13±0.118	
Cl ⁻	HSL	102.48±2.435	101.12±2.782	<0.0001
	HES	101.79±6.077	104.44±3.101	
Mg ⁺⁺	HSL	1.882±0.228	1.952±0.352	0.130
	HES	1.904±0.154	1.881±0.139	
Glucose	HSL	185.92±54.489	169.04±65.398	0.085
	HES	187.98±58.001	158.19±52.291	
Lactate ⁻	HSL	1.457±0.806	6.098±2.010	<0.0001
	HES	1.356±0.469	1.748±0.699	
Hb	HSL	12.436±1.692	11.806±1.634	0.409
	HES	12.735±1.662	11.956±1.291	
Ht	HSL	38.041±5.164	36.062±4.987	0.344
	HES	39.133±5.255	36.386±3.865	
PO ₂	HSL	231.760±64.253	219.140±66.814	0.359
	HES	245.917±69.838	219.417±38.828	
PCO ₂	HSL	32.940±5.223	28.660±5.278	0.673
	HES	33.083±4.885	29.223±4.339	
pH	HSL	7.44±0.06	7.52±0.06	<0.0001
	HES	7.44±0.04	7.48±0.05	
BE	HSL	-1.006±2.422	1.338±2.268	<0.0001
	HES	-0.925±1.664	-1.250±1.968	
SaO ₂	HSL	96.564±1.027	96.352±1.062	0.385
	HES	96.529±0.798	96.577±1.484	
HCO ₃	HSL	22.080±2.355	23.264±2.463	<0.0001
	HES	22.202±1.857	21.179±1.907	

Sodium (Na⁺); Potassium (K⁺); Chloride (Cl⁻); Calcium (Ca⁺⁺); Magnesium (Mg⁺⁺); Glucose, lactate and bicarbonate (HCO₃) are in mmol/L; Hemoglobin (Hb) is in g/L; Haematocrit (Ht) and SaO₂ are in %, PaO₂ and PaCO₂ are in torr. Data are expressed as mean±SD. Statistical comparison was carried out with Student *t* test. *The difference was significant between HSL and HES groups. HSL – Hyperosmolar sodium lactate; HES – Hydroxyethyl starch

Table 5: Fluid parameters

Variables (mL)	HSL n=50	HES n=48	P value
Total fluid intake	2409.16±656.328	2652.229±786.408	0.099
Crystalloid	1058.00±374.896	1255.417±476.003	0.024
Colloid	694.783±399.305	1000.00±377.965	0.069
PC	289.229±150.727	291.267±131.207	0.954
TC	123.158±31.632	131.250±68.099	0.635
FFP	270.682±75.395	271.053±94.262	0.984
Total fluid output	2855.10±879.393	2543.75±1036.53	0.112
Total urine output	1679.00±647.92	1442.71±709.04	0.088
Total bleeding	1176.00±896.378	1101.04±804.475	0.664
Fluid balance	-445.94±815.305	108.479±1219.909	0.009

HSL – Hyperosmolar sodium lactate; HES – Hydroxyethyl starch; PC – Packed red cell; TC – Thrombocyte concentrate; FFP – Fresh frozen plasma; SD – Standard deviation. Data is expressed as mean±SD. P values are obtained from the independent *t* test with the significant different if P<0.05

DISCUSSION

Maximizing the CO by perioperative intravascular expansion during cardiac surgery is associated with a better improvement of tissue perfusion, reduced post-operative morbidity and length of stay.^[1-3] Patients undergoing cardiac surgery may not tolerate large volumes of fluid required for haemodynamic stabilisation due to reduction of cardiac performance. Hence, small volume infusion with an adequate intravascular expansion effect and ability to remove extracellular fluid excess may benefit these patients.^[4,5] Colloids have been widely used in perioperative cardiac surgery and were associated with lower net post-operative weight gain,^[4,6,7] however, high cost and potential risk of colloid usage are still unsolved issues. Hyperosmolar solution infusion has been shown to be beneficial during cardiac surgery.^[4,8-12] A study in cardiac surgery comparing colloid with hypertonic saline infusion found that excretion of excess body fluid was higher in hypertonic saline group than colloid and avoiding tissue oedema.^[13]

In this study, we used HES 6% as a reference solution because this solution has been widely used during perioperative cardiac surgery.^[4,22] Similar volumes of hypertonic sodium lactate and HES 6% was administered during cardiac surgery. Lactate plays an important role as an energy substrate for cardiac cells and many studies showed that lactate administration directly improved cardiac performance.^[16-21] Previous studies have showed that administration of HSL for post cardiac surgery patients resulted in better cardiac performance and tissue perfusion when compared with hypertonic sodium chloride 3%.^[14] Until now, we did not find any publication comparing the efficacy of HSL with colloid in cardiac surgery setting, therefore we conducted this study. The randomisation resulted in comparable baseline parameters between HSL and HES group.

This study revealed that HSL infusion resulted higher increase of CI with lower SVRI than HES infusion despite comparable intravascular volume expansion, as indicated by comparable changes in MAP and CVP in both groups. Multivariate analysis showed that the decrease of SVRI was mostly associated with the improvement of CI and this finding was consistent with the previous studies.^[10-15] The decrease of PVR/PVRI and also alkalinizing effect of HSL also contributed to improved cardiac performance. The increase in

Table 6: Utilization of concomitant drugs

Concomitant drugs	HSL Patients (%)	HES Patients (%)	P value
Dobutamin	15 (30.0)	21 (43.7)	>0.05
Milrinone	1 (2.0)	3 (6.2)	>0.05
Dobutamin and milrinone	1 (2.0)	0 (0.0)	>0.05
Nitroglycerine and norepinephrin	0 (0.0)	3 (6.2)	>0.05
Nitroglycerine and amiodaron	1 (2.0)	2 (4.2)	>0.05

Treatment with inotropes or vasodilators during surgery in the two groups (HSL or HES). Values are number of patients (percent); HSL – Hyperosmolar sodium lactate; HES – Hydroxyethyl starch

myocardial contractility directly related to the increase in cardiac performance and lactate has been shown to improve cardiac contractility.^[14-18] Unfortunately in this study, we did not record the cardiac contractility after study fluid infusion, but we strongly feel that the lactate contributed for the improvement of cardiac performance. The decrease of vascular resistance after hypertonic solution infusion has been attributed to its hypertonicity,^[5,11,12] and we postulate that the same mechanism worked during HSL infusion in this study. Previous studies have showed that the intravascular volume expansion after hyperosmolar solution infusion as indicated by the improvement of MAP and CVP contributed for increasing CI^[11,12,23] however the vascular expansion was comparable between HSL and HES group in this study.

In this study, lower total fluid input with comparable total fluid output resulted in a much negative fluid balance in HSL group, whereas it became positive in HES group. This finding is in accordance with previous trials.^[5,13,15] This effect is could be due to a redistribution of interstitial fluid and mobilisation of intracellular fluid from swollen endothelial cells to intravascular space. Endothelial cell swelling and accumulation of interstitial fluid frequently occurred in major surgery patients.^[3,24-27] Reduction of endothelial cell swelling and mobilisation of extra vascular fluid into intravascular space following hypertonic solution administration were beneficial during cardiac surgery because it would reduce tissue oedema and improve microcirculation.^[4,5,14,26,27]

Overall, HSL could be considered as an ideal fluid in cardiac surgery due to its ability to maintain haemodynamic status and improve CI with negative fluid balance. The goal of perioperative fluid therapy is not only to maintain the effective circulatory volume, but also to avoid fluid overload whenever possible. Tissue oedema in surgical patients should be minimised because it increases the morbidity and mortality.^[26,27]

The infusion of HSL was well-tolerated since there was no adverse event during this study and laboratory parameters were still in normal range in both groups. Sodium level increased and chloride level decreased after HSL infusion, however the average levels were still within the normal limits. In contrast, in HES group sodium level slightly decreased and chloride levels increased. Hypernatremia, one of the potential side-effects of hyperosmolar fluid infusion, was not found in this study, as also was the case in previous studies.^[14,15] Osmolality and electroneutrality principles were suspected responsible for the decrease of plasma chloride level after sodium lactate infusion.^[28]

After study drugs administration, arterial pH, HCO₃ and base excess reached normal levels in HSL group, whereas in HES group base excess and HCO₃ were almost unchanged and still below normal range. This finding indicated that HSL infusion could completely reverse tissue acidosis, whereas HES infusion did not improve the existing tissue acidosis. Reversal of tissue acidosis in this study could be due to mild alkalisation effect of HSL. This was an expected result, considering that infusion of HSL will increase the strong ion difference as a consequence of higher sodium and lower chloride in the HL patients after lactate gets metabolised.^[14,15]

Number of patients requiring concomitant supportive drug during cardiac surgery such as vasodilator and inotropes was numerically less in HSL group. However the difference was not significant as and the sample size might not be adequate in this respect.

Limitations of this study

This study had some limitations including lack of double-blinding as well as the absence of the data expressing cardiac contractility after HSL or HES infusion. Another limitation was that for measurement of the effect of study drugs on haemodynamic status, measurement of the extra vascular lung water also is desirable and could not be measured due to lack of suitable monitors

CONCLUSION

This study demonstrated that HSL administration was safe and resulted in better improvement of CI and much lower vascular resistance with negative intra-operative fluid balance during CABG surgery when compared with HES infusion.

REFERENCES

- Gan TJ, Soppitt A, Maroof M, el-Moalem H, Robertson KM, Moretti E, *et al.* Goal-directed intraoperative fluid administration reduces length of hospital stay after major surgery. *Anesthesiology* 2002;97:820-6.
- Mythen MG, Webb AR. Perioperative plasma volume expansion reduces the incidence of gut mucosal hypoperfusion during cardiac surgery. *Arch Surg* 1995;130:423-9.
- Rosenthal MH. Intraoperative fluid management – What and how much? *Chest* 1999;115:106S-12.
- Boldt J. Volume therapy in cardiac surgery. *Ann Card Anaesth* 2005;8:104-16.
- Johnston WE. PRO: Fluid restriction in cardiac patients for noncardiac surgery is beneficial. *Anesth Analg* 2006;102:340-3.
- Jansen PG, te Velhuis H, Wildevuur WR, Huybregts MA, Bulder ER, van der Spoel HI, *et al.* Cardiopulmonary bypass with modified fluid gelatin and heparin-coated circuits. *Br J Anaesth* 1996;76:13-9.
- McIlroy DR, Kharasch ED. Acute intravascular volume expansion with rapidly administered crystalloid or colloid in the setting of moderate hypovolemia. *Anesth Analg* 2003;96:1572-7.
- Boldt J, Zickmann B, Thiel A, Herold C, Dapper F, Hempelmann G. Hyperosmolar volume replacement in heart surgery. *Anaesthesist* 1990;39:412-9.
- Boldt J, Zickmann B, Ballesteros M, Herold C, Dapper F, Hempelmann G. Cardiorespiratory responses to hypertonic saline solution in cardiac operations. *Ann Thorac Surg* 1991;51:610-5.
- Boldt J, Zickmann B, Herold C, Ballesteros M, Dapper F, Hempelmann G. Influence of hypertonic volume replacement on the microcirculation in cardiac surgery. *Br J Anaesth* 1991;67:595-602.
- Järvelä K, Koskinen M, Kaukinen S, Kööbi T. Effects of hypertonic saline (7.5%) on extracellular fluid volumes compared with normal saline (0.9%) and 6% hydroxyethyl starch after aortocoronary bypass graft surgery. *J Cardiothorac Vasc Anesth* 2001;15:210-5.
- Järvelä K, Kaukinen S. Hypertonic saline (7.5%) after coronary artery bypass grafting. *Eur J Anaesthesiol* 2001;18:100-7.
- Järvelä K, Kaukinen S. Hypertonic saline (7.5%) decreases perioperative weight gain following cardiac surgery. *J Cardiothorac Vasc Anesth* 2002;16:43-6.
- Mustafa I, Lerverve XM. Metabolic and hemodynamic effects of hypertonic solutions: Sodium-lactate versus sodium chloride infusion in postoperative patients. *Shock* 2002;18:306-10.
- Lerverve XM, Boon C, Hakim T, Anwar M, Siregar E, Mustafa I. Half-molar sodium-lactate solution has a beneficial effect in patients after coronary artery bypass grafting. *Intensive Care Med* 2008;34:1796-803.
- Kline JA, Thornton LR, Lopaschuk GD, Barbee RW, Watts JA. Lactate improves cardiac efficiency after hemorrhagic shock. *Shock* 2000;14:215-21.
- Barbee RW, Kline JA, Watts JA. Depletion of lactate by dichloroacetate reduces cardiac efficiency after hemorrhagic shock. *Shock* 2000;14:208-14.
- Levy B, Mansart A, Montemont C, Gibot S, Mallie JP, Regnault V, *et al.* Myocardial lactate deprivation is associated with decreased cardiovascular performance, decreased myocardial energetics, and early death in endotoxic shock. *Intensive Care Med* 2007;33:495-502.
- Gladden LB. Lactate metabolism: A new paradigm for the third millennium. *J Physiol* 2004;558:5-30.
- Mustafa I, Roth H, Hanafiah A, Hakim T, Anwar M, Siregar E, *et al.* Effect of cardiopulmonary bypass on lactate metabolism. *Intensive Care Med* 2003;29:1279-85.
- Himpe D, Neels H, De Hert S, Van Cauwelaert P. Adding lactate to the prime solution during hypothermic cardiopulmonary bypass: A quantitative acid-base analysis. *Br J Anaesth* 2003;90:440-5.
- Treib J, Baron JF, Grauer MT, Strauss RG. An international view of hydroxyethyl starches. *Intensive Care Med* 1999;25:258-68.
- Rocha-e-Silva M, Poli de Figueiredo LF. Small volume hypertonic resuscitation of circulatory shock. *Clinics (Sao Paulo)* 2005;60:159-72.
- Shires T, Williams J, Brown F. Acute change in extracellular fluids associated with major surgical procedures. *Ann Surg* 1961;154:803-10.
- Kirby RR. Perioperative fluid therapy and postoperative pulmonary edema: Cause-effect relationship? *Chest* 1999;115:1224-6.
- Lobo DN, Macafee DA, Allison SP. How perioperative fluid balance influences postoperative outcomes. *Best Pract Res Clin Anaesthesiol* 2006;20:439-55.
- Holte K, Sharrock NE, Kehlet H. Pathophysiology and clinical implications of perioperative fluid excess. *Br J Anaesth* 2002;89:622-32.
- Shackford SR, Norton CH, Todd MM. Renal, cerebral, and pulmonary effects of hypertonic resuscitation in a porcine model of hemorrhagic shock. *Surgery* 1988;104:553-60.

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Announcement

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The Indian College of Anaesthesiologists is an Academic body of the Indian Society of Anaesthesiologists. The ICA is registered as a Trust in New Delhi and functions under ISA through a MOU. Membership of the college is limited to ISA Members only. Membership fee Rs. 5,000/-. I request all members of ISA to become part of ICA.

For details contact: **Dr. B Radhakrishnan, CEO, ICA**

Email: ceoica@isaweb.in, brk_tvm@yahoo.com Mobile: +91 98470 63190

Dr. M V Bhimeshwar
Hon. Secretary - ISA