

Effect of baseline serum albumin concentration on outcome of resuscitation with albumin or saline in patients in intensive care units: analysis of data from the saline versus albumin fluid evaluation (SAFE) study

Saline versus Albumin Fluid Evaluation Study Investigators

Abstract

Objective To determine whether outcomes of resuscitation with albumin or saline in the intensive care unit depend on patients' baseline serum albumin concentration.

Design Analysis of data from a double blind, randomised controlled trial.

Setting Intensive care units of 16 hospitals in Australia and New Zealand.

Participants 6045 participants in the saline versus albumin fluid evaluation (SAFE) study.

Interventions Fluid resuscitation with 4% albumin or saline in patients with a baseline serum albumin concentration of 25 g/l or less or more than 25 g/l.

Main outcome measures Primary outcome was all cause mortality at 28 days. Secondary outcomes were length of stay in the intensive care unit, length of stay in hospital, duration of renal replacement therapy, and duration of mechanical ventilation.

Main results The odds ratios for death for albumin compared with saline for patients with a baseline serum albumin concentration of 25 g/l or less and more than 25 g/l were 0.87 and 1.09, respectively (ratio of odds ratios 0.80, 95% confidence interval 0.63 to 1.02); $P = 0.08$ for heterogeneity. No significant interaction was found between baseline serum albumin concentration as a continuous variable and the effect of albumin and saline on mortality. No consistent interaction was found between baseline serum albumin concentration and treatment effects on length of stay in the intensive care unit, length of hospital stay, duration of renal replacement therapy, or duration of mechanical ventilation.

Conclusion The outcomes of resuscitation with albumin and saline are similar irrespective of patients' baseline serum albumin concentration.

Trial registration ISRCTN76588266.

Introduction

Intravenous fluid is fundamental to the management of patients in intensive care units. The two broad categories of fluid available are colloids and crystalloids. Among colloids, human albumin is unique in being a major human plasma protein that has important physiological roles.¹⁻⁴ As hypoalbuminaemia is common in acute illness and is associated with an increased risk of death,⁵ the use of albumin to simultaneously treat hypoalbuminaemia and

to increase intravascular volume seems intuitively attractive. A meta-analysis by the 1998 Cochrane Albumin Reviewers, however, suggested that giving albumin to critically ill patients for the treatment of both hypovolaemia and hypoalbuminaemia increased the risk of mortality.⁶ Subsequently the saline versus albumin fluid evaluation (SAFE) study reported no important difference in the overall risk of death for adults given albumin or saline for intravascular fluid resuscitation in intensive care units.⁷⁻⁸ In an updated meta-analysis incorporating data from the saline versus albumin fluid evaluation study, the Cochrane Injuries Group Albumin Reviewers concluded that there is no evidence that albumin reduces the risk of mortality in critically ill patients but a suggestion that it may increase the risk of death in patients with hypoalbuminaemia and burns.⁹ Thus while the saline versus albumin fluid evaluation study provides greater certainty over the effect of resuscitation with albumin or saline in a heterogeneous population of patients in intensive care units, effects in more selected populations of critically ill patients remain unknown. Using data from the saline versus albumin fluid evaluation study, we determined whether outcomes are influenced by baseline serum albumin concentration and whether either fluid can be recommended on the basis of patients' baseline serum albumin concentration.

Methods

Details of the saline versus albumin fluid evaluation study have been published elsewhere.⁷⁻⁸ The double blind, randomised controlled trial was carried out in the multidisciplinary intensive care units of 16 hospitals in Australia and New Zealand between November 2001 and June 2003.

Eligible adults were randomly assigned to receive either 4% albumin (Albumex; CSL, Melbourne, Australia) or normal saline for all fluid resuscitation in the intensive care unit until death, discharge, or 28 days after randomisation. Patients were excluded who had been admitted to the intensive care unit after cardiac surgery or liver transplantation or for the treatment of burns.

The primary outcome was all cause mortality within 28 days of randomisation. Secondary outcomes were length of stay in the intensive care unit, length of stay in hospital, duration of mechanical ventilation, and duration of renal replacement therapy.

Statistical analysis

We used χ^2 tests for categorical variables and t tests or analysis of variance for continuous variables to assess the association of

baseline variables, including baseline albumin concentration, with mortality at 28 days. Baseline covariates were then fitted to logistic regression models to determine those independently associated with mortality.

We examined baseline albumin concentration as a binary variable using a predetermined cut-off (≤ 25 g/l or > 25 g/l), and as a continuous variable. We assessed the effect of treatment allocation and baseline albumin concentration on 28 day mortality using logistic regression; we used the interaction between baseline albumin concentration and treatment assignment to examine whether the risk of death for those assigned to albumin compared with those assigned to saline was consistent between different baseline albumin concentrations. Initially we carried out the logistic regression without adjustment for other baseline risk factors; then adjusted for those covariates significant at the $P < 0.10$ level. We excluded central venous pressure and urine output owing to missing values (3136 for central venous pressure and 871 for urine output). We also examined the heterogeneity of treatment effect on the secondary outcomes of the saline versus albumin fluid evaluation study.

Results

The treatment groups had similar baseline characteristics, published previously.⁸ Data on baseline serum albumin concentration were available for 6045 patients, 3014 of those assigned to albumin and 3031 assigned to saline. No difference was found in the distribution of baseline serum albumin concentration by treatment group ($P = 0.76$).

The distribution of baseline characteristics within each stratum of baseline albumin concentration (≤ 25 g/l or > 25 g/l) was similar in those assigned to albumin and those assigned to saline (table 1).

Overall 2451 (40.5%) patients had a baseline serum albumin concentration of 25 g/l or less (1228 patients (50.1%) in the albumin group and 1223 patients (49.9%) in the saline group). Compared with patients with a baseline serum albumin concentration of more than 25 g/l ($n = 3594$), those with a concentration of 25 g/l or less were older and were more likely to be admitted to the intensive care unit after surgery, more likely to have severe sepsis or acute respiratory distress syndrome, and less likely to have had traumatic brain injury. Severity of illness, as assessed by the acute physiology and chronic health evaluation II score,¹⁰ was similar in patients with a baseline serum albumin concentration of 25 g/l or less or more than 25 g/l (table 2).

The primary outcome measure (mortality at 28 days) and baseline serum albumin concentration were available for 6040 (99.9%) patients.

Baseline serum albumin concentration as both a binary and a continuous variable was independently associated with mortality. Other baseline factors independently associated with mortality were age, reason for admission to the intensive care unit, acute physiology and chronic health evaluation II score, liver and cardiovascular components of the sequential organ failure assessment score,¹¹ mechanical ventilation at baseline, and heart rate ($P < 0.10$).

After adjusting for baseline risk factors for death, a baseline serum albumin concentration of 25 g/l or less was independently associated with risk of death (odds ratio 1.30, 95% confidence interval 1.16 to 1.51, $P = 0.0009$). The findings were similar (table 3) when baseline serum albumin concentration was treated as a continuous variable (odds per 1 g/l decrease in baseline serum albumin concentration 1.02, 95% confidence interval 1.01 to 1.03, $P < 0.0001$).

Table 1 Baseline characteristics of patients in intensive care units assigned to albumin or saline, stratified by baseline serum albumin concentrations (≤ 25 g/l or > 25 g/l). Values are numbers (percentages) of patients unless stated otherwise

Characteristic	Albumin group	Saline group
Baseline serum albumin concentration ≤ 25 g/l*		
Mean (SD) age (years)	61.5 (18.4)	61.1 (17.7)
Men	725 (59.0)	715 (58.5)
Admitted to intensive care unit for postoperative care	679 (55.3)	654 (53.5)
Present at baseline:		
Traumatic brain injury	52 (4.2)	46 (3.8)
Severe sepsis	290 (24.1)	314 (26.5)
Acute respiratory distress syndrome	33 (2.7)	45 (3.7)
Mean (SD) acute physiology and chronic health evaluation II score	19.0 (7.6)	19.1 (7.9)
Mean (SD) urine output (ml/h)	85.0 (114.9)	91.9 (174.0)
Mean (SD) mean arterial pressure (mm Hg)	75.8 (15.3)	76.9 (15.7)
Receiving renal replacement therapy	23 (1.9)	23 (1.9)
Receiving mechanical ventilation	744 (60.7)	762 (62.4)
Prior treatment with angiotensin converting enzyme inhibitor	163 (13.3)	159 (13.1)
Baseline serum albumin concentration > 25 g/l†		
Mean (SD) age (years)	56.0 (19.3)	56.1 (19.0)
Men	1064 (59.6)	1124 (62.2)
Admitted to intensive care unit for postoperative care	576 (32.3)	599 (33.1)
Present at baseline:		
Traumatic brain injury	172 (9.6)	190 (10.5)
Severe sepsis	251 (14.4)	254 (14.4)
Acute respiratory distress syndrome	19 (1.1)	18 (1.0)
Mean (SD) acute physiology and chronic health evaluation II score	19.0 (8.0)	19.4 (8.1)
Mean (SD) urine output (ml/h)	92.4 (140.5)	97.1 (157.8)
Mean (SD) mean arterial pressure (mm Hg)	79.7 (16.8)	79.5 (16.2)
Receiving renal replacement therapy	19 (1.1)	16 (0.9)
Receiving mechanical ventilation	1225 (68.6)	1230 (68.0)
Prior treatment with angiotensin converting enzyme inhibitor	249 (13.9)	268 (14.9)

*1228 participants in albumin group, 1223 in saline group.

†1786 participants in albumin group, 1808 in saline group.

Patients assigned to albumin had a higher mean serum albumin concentration during the first seven days after randomisation (fig 1). This difference was apparent for patients with a baseline serum albumin concentration of 25 g/l or less or more than 25 g/l ($P < 0.0001$ for both). On average, patients assigned to albumin received a lower daily volume of resuscitation fluid than patients assigned to saline (table 4).

Patient outcomes

Among patients with a serum albumin concentration of 25 g/l or less, deaths occurred in 291 (23.7%) assigned to albumin and 321 (26.2%) assigned to saline (odds ratio 0.87, 95% confidence interval 0.73 to 1.05, $P = 0.14$). In patients with a serum albumin concentration of more than 25 g/l, deaths occurred in 353 (19.8%) assigned to albumin and 334 (18.5%) assigned to saline (odds ratio 1.09, 95% confidence interval 0.92 to 1.28, $P = 0.33$; fig 2 and table 5). The ratio of the odds ratios for patients with a baseline serum albumin concentration of 25 g/l or less and of more than 25 g/l was 0.80 (95% confidence interval 0.63 to 1.02). After adjustment for baseline risk factors for death (table 3), the odds ratios for death of patients with a baseline serum albumin concentration of 25 g/l or less and more than 25 g/l given albumin compared with saline were 0.84 and 1.15, respectively (ratio of odds ratios 0.73, 95% confidence interval 0.55 to 0.97). The P

Table 2 Comparison of baseline characteristics of patients in saline versus albumin fluid evaluation study with baseline serum albumin concentration 25 g/l or less or more than 25 g/l. Values are numbers (percentages) of patients unless stated otherwise

Characteristic	Baseline serum albumin concentration \leq 25 g/l (n=2451)	Baseline serum albumin concentration >25 g/l (n=3594)	P value
Mean (SD) age (years)	61.3 (18.0)	56.1 (19.1)	<0.0001
Men	1440 (58.8)	2188 (60.9)	0.10
Admitted to intensive care unit for postoperative care	1333 (54.4)	1175 (32.7)	<0.0001
Present at baseline:			
Traumatic brain injury	98 (4.0)	362 (10.1)	<0.0001
Severe sepsis	604 (24.6)	505 (14.1)	<0.0001
Acute respiratory distress syndrome	78 (3.2)	37 (1.0)	<0.0001
Mean (SD) acute physiology and chronic health evaluation II score	19.1 (7.8)	19.2 (8.1)	0.44
Mean (SD) urine output (ml/h)	88.4 (146.9)	94.8 (149.5)	0.15
Mean (SD) mean arterial pressure (mm Hg)	76.4 (15.5)	79.6 (16.5)	<0.0001
Receiving renal replacement therapy	46 (1.9)	35 (1.0)	0.003
Receiving mechanical ventilation	1506 (61.4)	2455 (68.3)	<0.0001
Prior treatment with angiotensin converting enzyme inhibitor	322 (13.1)	517 (14.4)	0.18

values for heterogeneity of treatment effects between those with baseline serum albumin concentration of 25 g/l or less and of more than 25 g/l with and without adjustment for baseline risk

Table 3 Multivariate analysis of association of baseline characteristics (including baseline serum albumin concentration) and assigned group (albumin or saline) with risk of death

Baseline characteristic	P value	Odds ratio* (95% CI)
Albumin as binary variable:		
Age	<0.0001	1.030 (1.025 to 1.035)
Source of admission to intensive care unit	<0.0001	
Acute physiology and chronic health evaluation II score	<0.0001	1.072 (1.061 to 1.083)
Sequential organ failure assessment score:		
Liver component	<0.0001	—
Cardiovascular component	<0.0001	—
Mechanical ventilation	0.001	1.326 (1.126 to 1.570)
Heart rate	<0.0001	1.007 (1.003 to 1.010)
Baseline serum albumin concentration as binary variable	0.0009	0.772 (0.663 to 0.899)
Albumin v saline group	0.999	1.000 (0.869 to 1.151)
Albumin as continuous variable:		
Age	<0.0001	1.030 (1.025 to 1.034)
Source of admission to intensive care unit	<0.0001	
Acute physiology and chronic health evaluation II score	<0.0001	1.072 (1.061 to 1.083)
Sequential organ failure assessment score:		
Liver component	<0.0001	—
Cardiovascular component	<0.0001	—
Mechanical ventilation	0.0008	1.334 (1.126 to 1.580)
Heart rate	0.001	1.006 (1.003 to 1.010)
Baseline serum albumin concentration as continuous variable	<0.0001	0.979 (0.969 to 0.989)
Albumin v saline group	0.944	0.995 (0.864 to 1.145)

*For age, per one year increase in age; for acute physiology and chronic health evaluation II score, per one point increase in score; for heart rate, per one beat per minute increase in rate.

factors (P values not adjusted for multiple comparisons) were 0.08 and 0.04, respectively (fig 2 and table 5). When baseline serum albumin concentration was treated as a continuous variable, no significant interaction was found between baseline concentration and treatment assignment for 28 day mortality (unadjusted P = 0.73, adjusted P = 0.94). No significant interaction was found between baseline serum albumin concentration and treatment group for length of stay in the intensive care unit, duration of mechanical ventilation, and duration of renal replacement therapy (P values 0.50, 0.85, and 0.33, respectively), but an interaction of borderline significance was found for length of hospital stay (P = 0.05 without correction for multiple hypothesis testing; table 5).

Discussion

Our study does not provide evidence that the effect of resuscitation with albumin compared with saline in the intensive care unit is different in patients with different baseline serum albumin concentrations. Nor does it provide evidence to support the suggestion that albumin increases the risk of mortality in patients with hypoalbuminaemia. When the odds ratios for death was compared in patients with a baseline serum albumin concentration of 25 g/l or less or of more than 25 g/l we found only limited evidence that treatment effects were different and this only after correction for other baseline risk factors. When we considered the effect of baseline serum albumin concentration as a continuous variable across the spectrum of albumin concentrations, baseline concentration had no impact on the treatment effect even after correction for other baseline risk factors. Taken together these results suggest that albumin and saline produce similar treatment effects across the range of albumin concentrations observed in our study.

A recent meta-analysis assessed the effect of resuscitation with albumin or other fluids on outcomes other than mortality. The authors included 71 trials, which randomised 3782 acutely ill adults, children, and neonatal patients and concluded that albumin significantly reduced overall morbidity.¹² This contrasts with the findings of the saline versus albumin fluid evaluation study in which there were no significant differences in morbidity in patients resuscitated with either albumin or saline.⁸ In the current analysis we found no interaction between baseline serum albumin concentration and choice of resuscitation fluid for length of stay in the intensive care unit, duration of renal replacement therapy, or duration of mechanical ventilation. We found borderline evidence of an interaction for length of hospital stay; patients with a baseline serum albumin concentration more than 25 g/l who were resuscitated with albumin had a shorter hospital stay (mean difference 0.6 days). In the absence of any other significant heterogeneity, we interpret this as a chance finding.

The strengths of our study are the size and methodological strengths of the saline versus albumin fluid evaluation study, a double blind randomised controlled trial that has been recognised as a high quality landmark study and which contributed 82.8% of the data for the latest Cochrane meta-analysis.^{9 13} The limitations are the inherent weaknesses of subgroup analyses and that the saline versus albumin fluid evaluation study was not primarily designed to determine the effect of treating hypoalbuminaemia with albumin. In the saline versus albumin fluid evaluation study, patients received the amount of fluid the treating clinician thought necessary to restore or maintain intravascular volume, and albumin was not given to achieve or maintain a particular serum albumin concentration. Patients assigned

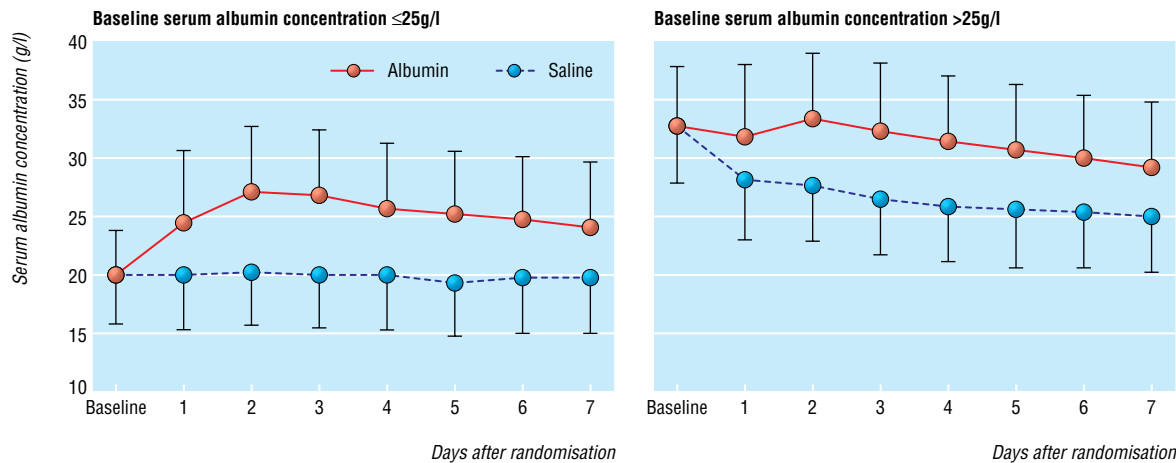


Fig 1 Effect of fluid resuscitation with albumin or saline on mean serum albumin concentration for seven days after randomisation stratified by baseline serum albumin concentration (≤ 25 g/l or > 25 g/l). Error bars indicate standard deviation

to albumin maintained higher serum albumin concentrations but this was not associated with clear benefit. Administering albumin to achieve and maintain a greater increase in serum albumin concentration, for instance to more than 30 g/l, has been advocated⁸; whether this would improve patients outcomes requires an appropriately designed trial. Although recognising the limitations of our analysis, the 2004 Cochrane review reported only 637 patients and 86 deaths from 10 trials of patients treated for hypoalbuminaemia; six trials (451 patients) were in adults.⁹ Of the 6997 adults in the saline versus albumin fluid evaluation study, 2451 had a recorded baseline serum albumin concentration of 25 g/l or less of whom 612 died. Our analysis provides most of the available data on the effect of albumin administration to adults with hypoalbuminaemia in an

intensive care unit, and so may provide a more reliable estimate of the treatment effect of albumin than the data on which the 2004 Cochrane review is based.

In conclusion, resuscitation with albumin and saline in the intensive care unit produces similar patient outcomes irrespective of patients' baseline serum albumin concentrations. Although albumin does not increase the risk of mortality, evidence is insufficient to support the routine use of albumin to maintain or increase intravascular volume in adults with hypoalbuminaemia in intensive care units. Whether administering albumin to adults in intensive care units to maintain a particular serum albumin concentration would be beneficial could only be answered by an appropriately designed trial. Our

Table 4 Volume of albumin or saline administered per day for first seven days in intensive care by treatment group and baseline serum albumin concentration

Day	Baseline serum albumin concentration ≤ 25 g/l				P value (albumin v saline)	Baseline serum albumin concentration > 25 g/l				P value (albumin v saline)
	Mean (SD) albumin administered per day (ml)		Mean (SD) saline administered per day (ml)			Mean (SD) albumin administered per day (ml)		Mean (SD) saline administered per day (ml)		
	No*		No*			No*		No*		
1	1221	1217 (981)	1219	1650 (1585)	<0.0001	1777	1154 (970)	1808	1470 (1469)	<0.001
2	1093	648 (973)	1104	1211 (1745)	<0.0001	1589	550 (803)	1621	752 (1208)	<0.0001
3	797	281 (554)	830	429 (911)	<0.0001	1154	259 (554)	1158	290 (617)	0.21
4	640	221 (449)	642	312 (846)	0.02	859	179 (420)	870	171 (456)	0.71
5	521	190 (416)	517	278 (674)	0.01	708	183 (448)	723	173 (463)	0.68
6	447	183 (447)	444	238 (617)	0.13	593	164 (428)	599	151 (486)	0.61
7	390	197 (474)	390	241 (682)	0.30	514	172 (469)	514	131 (381)	0.13

Mean (standard deviations) rounded to nearest integer.

*Patients still in intensive care unit.

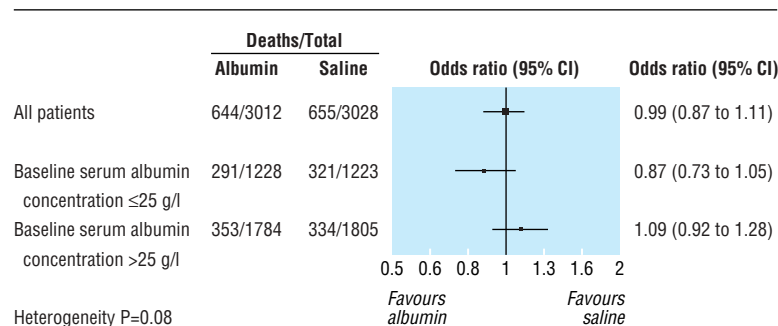


Fig 2 Unadjusted odds ratio (95% confidence interval) of death in all patients and in subgroups with baseline serum albumin concentration of 25 g/l or less and of more than 25 g/l. (Heterogeneity of treatment effect in subgroups with baseline serum albumin concentration ≤ 25 g/l v > 25 g/l, $P=0.08$)

Table 5 Primary and secondary outcomes stratified by baseline serum albumin concentration 25 g/l or less and more than 25 g/l. Values are means (standard deviations) unless stated otherwise

Outcome	Albumin group	Saline group	Odds ratio (95% CI)	Absolute difference (95% CI)	P value
Baseline serum albumin \leq25 g/l					
Status at 28 days:	n=1228	n=1223	—	—	—
No (%) dead	291 (23.7)	321 (26.2)	0.87 (0.73 to 1.05)	—	0.14
No (%) alive in intensive care	54 (4.4)	33 (2.7)	1.66 (1.07 to 2.58)	—	0.02
No (%) alive in hospital	333 (27.1)	322 (26.3)	1.04 (0.87 to 1.25)	—	0.66
Length of stay in intensive care unit (days)	6.9 (6.9)	6.5 (6.4)	—	0.32 (−0.21 to 0.84)	0.24
Length of hospital stay (days)	16.8 (9.5)	16.4 (9.5)	—	0.40 (−0.35 to 1.16)	0.30
Duration of mechanical ventilation (days)	4.8 (6.5)	4.6 (5.8)	—	0.22 (−0.27 to 0.71)	0.37
Duration of renal replacement therapy (days)	0.6 (2.4)	0.6 (2.4)	—	0.01 (−0.18 to 0.20)	0.89
Baseline serum albumin concentration $>$25 g/l					
Status at 28 days:	n=1784	n=1805	—	—	—
No in group	1784	1805	—	—	—
No (%) dead	353 (19.8)	334 (18.5)	1.10 (0.92 to 1.28)	—	0.33
No (%) alive in intensive care unit	39 (2.2)	40 (2.2)	0.99 (0.63 to 1.54)	—	0.95
No (%) alive in hospital	376 (21.1)	436 (24.2)	0.84 (0.73 to 0.98)	—	0.03
Length of stay in intensive care unit (days)	6.2 (6.3)	6.2 (6.2)	—	0.09 (−0.31 to 0.50)	0.65
Length of hospital stay (days)	14.4 (9.5)	15.0 (9.7)	—	−0.60 (−1.23 to 0.03)	0.06
Duration of mechanical ventilation (days)	4.4 (5.8)	4.3 (5.6)	—	0.16 (−0.21 to 0.53)	0.39
Duration of renal replacement therapy (days)	0.4 (2.1)	0.3 (1.6)	—	0.12 (−0.00 to 0.25)	0.05

Ratio of odds ratios for death (95% confidence interval for ratio) for albumin versus saline with baseline serum albumin concentration \leq 25 g/l v $>$ 25 g/l; uncorrected 0.80 (0.63 to 1.02), corrected 0.73 (0.55 to 0.97). P values for interaction terms (heterogeneity): (independent variables=treatment (albumin or saline) and baseline serum albumin (\leq 25 g/l or $>$ 25 g/l)). Mortality at 28 days: uncorrected P=0.08, corrected P=0.04. Length of stay in intensive care unit: uncorrected P=0.50. Length of stay in hospital: uncorrected P=0.05. Duration of mechanical ventilation: uncorrected P=0.85. Duration of renal replacement therapy: uncorrected P=0.33.

data suggest that any benefit (or harm) would be small and so a very large trial would be required.

Contributors: Investigators for the saline versus albumin fluid evaluation study are: writing committee—Simon Finfer (chair), Rinaldo Bellomo, Suzanne McEvoy, Sing Kai Lo, John Myburgh, Bruce Neal, and Robyn Norton; management committee—Robyn Norton (chair), Julie French (senior project manager), Rinaldo Bellomo, Simon Finfer, John Myburgh, Suzanne McEvoy, Gordon Doig, Mary Hayek, and Sheridan O'Donnell; steering committee—Simon Finfer (chair), Anthony Bell, Rinaldo Bellomo, Neil Boyce, David Blythe, John Cade, Marianne Chapman, Louise Cole, James Cooper, Andrew Davies, Craig French, Julie French, Christopher Joyce, Colin McArthur, Stephen MacMahon, John Myburgh, Bruce Neal, Robyn Norton, Jeffrey Presneill, Peter Saul, Ian Seppelt, Dianne Stephens, Andrew Turner, Anthony Williams, and Clive Woolfe; and external safety and data monitoring committee—Richard Peto (chair), Peter Sandercock, Charles Sprung, and J Duncan Young. The statistical analysis was carried out by Sing Kai Lo, Siva Sivarajasingham, Lesley Francis, and Mark Woodward (George Institute for International Health, University of Sydney, NSW). The site investigators were: Julie Charlton, James Cooper, Andrew Davies, Catherine Harry, Lisa Higgins, Katherine Moulden, and Shirley Vallance (Alfred Hospital, Melbourne, Vic); Janine Chadderton, Lynette Newby, and Colin

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Competing interests: The saline versus albumin fluid evaluation study was part funded by CSL. CSL has acted as a sponsor for scientific meetings of the Australian and New Zealand Intensive Care Society and its clinical trials

What is already known on this topic

Administering albumin may increase the risk of death in critically ill patients with hypoalbuminaemia

What this study adds

Irrespective of patients' baseline serum albumin concentration, fluid resuscitation with albumin or saline produced similar outcomes

Although albumin does not increase the risk of mortality in patients with hypoalbuminaemia, data do not support its routine use to maintain or increase intravascular volume in critically ill adults

group. CSL has paid travel expenses for Simon Finfer and Rinaldo Bellomo to present the results of the SAFE study at scientific and industry sponsored meetings. Andrew Davies and Diane Stephens own shares in CSL.

Ethical approval: This study was approved by the ethics committees of the University of Sydney and of each of the participating institutions.

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