

### 5.3 Strategies to Optimize Delivery and Minimize Risks of EN: Small Bowel Feeding vs. Gastric

**Question:** Does enteral feeding via the small bowel compared to gastric feeding result in better outcomes in the critically ill adult patient?

**Summary of evidence:** There were seventeen randomized trials that were reviewed, all of which were level 2 studies. In the Taylor et al study, only 34% of the patients achieved small bowel access in this study (large number of protocol violations) and hence the meta-analysis was done with and without this study. Minard et al compared outcomes in patients receiving early immune enhanced enteral nutrition via the small bowel to those receiving delayed immune enhanced enteral nutrition via the gastric route. Meta-analyses on mortality, infections & time dependent variables (LOS) were done with and without the Minard study.

**Mortality:** Based on the 14 studies that reported on mortality, no significant differences between the groups were found (RR 1.01, 95% CI 0.84, 1.22,  $p=0.89$ , heterogeneity  $I^2=0\%$ ; figure 1). When the Taylor et al & Minard studies was excluded, the relative risk did not change (RR 1.03, 95% CI 0.85, 1.24,  $p=0.77$ , heterogeneity  $I^2=0\%$ ; figure 2).

**Infections (Pneumonia):** Based on the 14 studies that reported on pneumonia, the meta-analysis showed that small bowel feeding was associated with a reduction in pneumonia when compared to gastric feeding (RR 0.78, 95% CI 0.61, 1.00,  $p=0.05$ , heterogeneity  $I^2=28\%$ ; figure 3). When the studies by Taylor et al and Minard et al were removed from the analysis, small bowel feeding was associated with only a trend in the reduction of pneumonia (RR 0.77, 95% CI 0.57, 1.06,  $p=0.11$ , heterogeneity  $I^2=36\%$ ; figure 4).

**LOS:** When all 10 studies that reported ICU LOS were aggregated, enteral feeding via the small bowel had no effect on ICU length of stay (WMD -1.19, 95% CI -3.46, 3.07,  $p=0.91$ , heterogeneity  $I^2=98\%$ ; figure 5). When the Minard study was excluded from the analysis, the signal did not change (WMD -0.86, 95% CI -4.25, 2.53,  $p=0.62$ , heterogeneity  $I^2=98\%$ ; figure 6). Based on the aggregation of the 5 studies that reported hospital LOS, enteral feeding via the small bowel had no effect on hospital length of stay (WMD 0.56, 95% CI -3.60, 4.73,  $p=0.79$ , heterogeneity  $I^2=24\%$ ; figure 7) when compared to gastric feeding.

**Ventilator days:** Based on the aggregation of the 6 studies that reported duration of ventilation, enteral feeding via the small bowel compared to gastric feeding had no effect on duration of ventilation (WMD -0.89, 95% CI -2.75, 0.97,  $p=0.35$ , heterogeneity  $I^2=81\%$ ; figure 8).

**Nutritional Outcomes:** Many studies reported on nutritional complications, such as GI bleeds, vomiting, diarrhea, constipation and abdominal bloating. There was no difference between the 2 groups in some studies (Davies 2011, White, Eatock, Friedman), while other reported a significant improvement in nutritional outcomes in the group fed via small bowel such as better nutrition efficiency (Hsu, Acosta-Escribano), calorie/protein intake & less time to reach goal (Hsu), vomiting (Hsu) and significantly less gastrointestinal tract colonization and high gastric residual volumes

(Acosta Escribano). The studies that reported nutritional delivery generally showed better success at meeting goal targets and reaching them sooner. However, this could also be explained by the confounded nature of different gastric feeding strategies. When the data from the 6 studies that reported nutritional efficiency (% goal rate received) as a mean  $\pm$  standard deviation were aggregated, small bowel feeding compared to gastric feeding was associated with a significantly greater percentage of nutritional efficiency (WMD 10.59, 95% CI 4.76, 16.41,  $p=0.0004$ , heterogeneity  $I^2=88\%$ ; figure 9). When the data from the 4 studies that reported the time to reach nutritional goal rate were aggregated, small bowel feeding compared to gastric feeding had no effect on the time to reach nutritional goals (WMD -3.41, 95% CI -13.45, 6.62,  $p=0.51$ , heterogeneity  $I^2=87\%$ ; figure 10). One study (Friedman 2015) reported a significant increase in cost when using small bowel vs gastric feeds, though the details on this calculation and the statistical significance was not reported.

**Other complications:** The group that had a more aggressive feeding regimen and small bowel feeding (Taylor) had fewer major complications and a better neurological outcome at 3 months than the group receiving gastric feeds.

**Conclusions:**

- 1) Small bowel feeding, compared to gastric feeding may be associated with a reduction in pneumonia in critically ill patients.
- 2) Small bowel feeding, compared to gastric feeding has no effect on mortality or ventilator days in critically ill patients receiving small bowel vs. gastric feedings.
- 3) Small bowel feeding is associated with improved calorie and protein intake and with less time taken to reach target rate of enteral nutrition when compared to gastric feeding.

*Level 1 study: if all of the following are fulfilled: concealed randomization, blinded outcome adjudication and an intention to treat analysis.*

*Level 2 study: If any one of the above characteristics are unfulfilled.*

**Table 1. Randomized studies evaluating small bowel feeding vs. gastric in critically ill patients**

Study	Population	Methods (score)	Mortality # (%)†		Pneumonia # (%)‡	
			Small bowel	Gastric	Small bowel	Gastric
<b>1. Montecalvo 1992</b>	Med/Surg ICU Anticipated feed >3days N=38 from 2 ICUs	C.Random: not sure ITT: no Blinding: no (8)	5/19 (26)	5/19 (26)	4/19 (21)	6/19 (32)
<b>2. Kortbeek 1999</b>	Trauma ISS>16 Vent >48h N=80 from 2 ICUs	C.Random: yes ITT: yes Blinding: no (11)	4/37 (11)	3/43 (7)	10/37 (27)	18/43 (42)
<b>3. Taylor 1999</b>	Head injured ventilated > 10 yrs N=82	C.Random: not sure ITT: yes Blinding: no (10)	<b>6-month</b> 5/41(12)	<b>6-month</b> 6/41 (15)	<b>Pneumonia</b> 18/41 (44)                      26/41 (63)  <b>Total Infections</b> 25/41 (61)                      35/41 (85)	
<b>4. Kearns 2000</b>	MICU Feed >3days APACHE -21 N=44	C.Random: not sure ITT: yes Blinding: no (9)	5/21 (24)	6/23 (26)	4/21 (19)	3/23 (13)
<b>5. Minard 2000</b>	Trauma GCS 3-10 N=27	C.Random: not sure ITT: no Blinding: no (7)	1/12 (8)	4/15 (27)	6/12 (50)	7/15 (47)
<b>6. Esparaza 2001</b>	MICU MV = 98% APACHE -25 N=54	C.Random: not sure ITT: yes Blinding: no (8)	10/27 (37)	11/27 (41)	NR	NR
<b>7. Boivin 2001</b>	Med/Surg/Neuro MV-98% Feed >72h APACHE-16 N=80	C.Random: not sure ITT: no Blinding: no (6)	18/39 (46)	18/39 (46)	NR	NR

<b>8. Day 2001</b>	Neurological ICU APACHE ~ 48 N=25	C.Random: not sure ITT: yes Blinding: no (5)	NR	NR	0/14 (0)	2/11 (18)
<b>9. Davies 2002</b>	Med/surg/trauma Feed > 3days MV=90%; APACHE~21 N=73	C.Random: not sure ITT: no Blinding no (8)	4/34 (12)	5/39 (13)	2/31 (6)	1/35 (3)
<b>10. Neumann 2002</b>	MICU N=60	C.Random: not sure ITT: yes Blinding: no (6)	NR	NR	NR	NR
<b>11. Montejo 2002</b>	14 ICU APACHE ~18 Feed >5days N=101 from 11 ICUs	C.Random: not sure ITT: yes Blinding: no (6)	19/50 (38)	22/51 (43)	16/50 (32)	20/51 (39)
<b>12. Hsu 2009</b>	Medical ICU Anticipated feed >3days N=121	C.Random: Yes ITT: Yes Blinding: No (9)	26/59 (44)	24/62 (39)	5/59 (9)	15/62 (24)
<b>13. White 2009</b>	Medical ICU mechanically ventilated >24hrs N=108	C.Random: Yes ITT: Yes Blinding: No (7)	11/50 (22)	5/54 (9)	5/50 (10)	11/54 (20)
<b>14. Acosta-Escribano 2010</b>	Traumatic brain injury, mechanically ventilated patients in ICU required EN for >5 days N=104	C.Random: No ITT: Yes Blinding: No (9)	<b>30-day</b> 6/50 (12)	<b>30-day</b> 9/54 (17)	16/50 (32)	31/54 (57)

15. Davies 2012	Critically ill , mechanically ventilated, on narcotic infusion with elevated GRV from 17 ICUs N=181	C.Random: Yes ITT: Yes Blinding: No (11)	13/91 (14)	12/89 (13)	18/91 (20)	19/89 (21)
16. Friedman 2015	Critically ill adults withour contraindication for enteral nutrition, expected ICU LOS >48 hrs N=115	C.Random: Yes ITT: Yes Blinding: No (9)	ICU 20/54 (37)	ICU 22/61 (36)	13/54 (24)	12/61 (20)
17. Wan 2015	Mixed ICU patients. Single Centre. N=70	C.Random: Yes ITT: Yes Blinding: No (8)	NR	NR	Aspiration pneumonia 0/35	Aspiration pneumonia 10/35

Table 1. Randomized studies evaluating small bowel feeding vs. gastric in critically ill patients (continued)

Study	LOS days		Ventilator days		Nutritional Outcomes		Other	
	Small bowel	Gastric	Small bowel	Gastric	Small bowel	Gastric	Small bowel	Gastric
1. Montecalvo 1992	ICU 11.7 ± 8.2 (19)	ICU 12.3 ± 10.8 (19)	10.2 ± 7.1 (19)	11.4 ± 10.8 (19)	Daily caloric intake (%) 61 ± 17	46.9 ± 25.9	GI bleeding 7/19 (37) Diarrhea 12/19 (63) Vomiting 3/19 (16)	GI bleeding 6/19 (32) Diarrhea 9/19 (47) Vomiting 3/19 (16)
2. Kortbeek 1999	ICU 10 (3-24) Hospital 30 (16-47)	ICU 7 (3-32) Hospital 25 (9-88)	9 (2-13)	5 (3-15)	Time to tolerate full feeds 34 ± 7.1	43.8 ± 22.6	NR	NR

3. Taylor 1999	NR	NR	NR	NR	<p>% energy needs met (mean) 59.2                      36.8</p> <p>% nitrogen needs met (mean) 68.7                      37.9</p>	<p>37 % major complications</p> <p>61 % had better neurological outcome at 3 months</p>	<p>61 % major complications</p> <p>39 % had better neurological outcome at 3months</p>
4. Kearns 2000	<p>ICU 17 ± 2 (21) Hospital 39 ± 10 (21)</p>	<p>ICU 16 ± 2 (23) Hospital 43 ± 11 (23)</p>	NR	NR	<p>Calories (kcal/kg/day) 18 ± 1                      12 ± 2</p> <p>Protein (gm/kg/day) 0.7 ± 0.1                      0.4 ± 0.1</p> <p>% REE delivered 69 ± 7                      47 ± 7</p>	Diarrhea 3 days	Diarrhea 2 days
5. Minard 2000	<p>ICU 18.5 ± 8.8 (12) Hospital 30 ± 14.7 (12)</p>	<p>ICU 11.3 ± 6.1 (12) Hospital 21.3 ± 14.7 (12)</p>	15.1 ± 7.5 (12)	10.4 ± 6.1 (15)	<p>Time feeding initiated (hours) 33 ± 15                      84 ± 41</p> <p>Avg kcals/ day 1509 ± 45                      1174 ± 425</p> <p>Days fed 13 ± 3.7                      8 ± 4.5</p> <p># patients with &gt; 50 % goal for ≥ 5 days 10/12 (83)                      7/15 (47)</p>	<p>Diarrhea 11/12 (92) Vomiting 1/12 (8)</p>	<p>Diarrhea 8/15 (53) Vomiting 3/15 (20)</p>
6. Esparaza 2001	NR	NR	NR	NR	<p>Feed days (average) 3.6                      4.1</p> <p>Average daily % of goal 66                      64</p>	NR	NR
7. Boivin 2001	NR	NR	NR	NR	<p>Time of placement 304 minutes                      13 minutes</p> <p>Time to goal rate achieved and maintained for 4 hours 33 hours                      32 hours</p>	NR	NR
8. Day 2001	NR	NR	NR	NR	<p>Calories and protein received were significantly higher only on days 2 and 3 in the gastric group. No difference between the groups on Days 1, 4-10.</p> <p>Replaced tubes 16/14                      9/11</p>	Diarrhea 7/14 (50)	Diarrhea 5/11 (45)

9. Davies 2002	ICU 13.9 ± 1.8 (34)	ICU 10.4 ± 1.2 (39)	NR	NR	Time to reach target rate 23.2 ± 3.9      23.0 ± 3.4 Time to start feeds 81.2 ± 13.4      54.5 ± 4.9	GI bleeding 3/31 (10) Diarrhea 4/31 (13)	GI bleeding 0/35 (0) Diarrhea 3/35 (9)
10. Neumann 2002	NR	NR	NR	NR	Time from initial attempt to start of feeding 27.0 ± 22.6      11.2 ± 11.0 Time to reach goal rate (from initial placement attempt) 43 ± 24.1      28.8 ± 15.9 Time to reach goal rate (from successful tube placement) 17.3 ± 15.7      17.0 ± 11.9	Aspiration 1/30 (3)	Aspiration 0/30 (0)
11. Montejo 2002	ICU 15 ± 10 (50)	ICU 18 ± 16 (50)	NR	NR	High gastric residuals 1/50 (2)      25/51 (49) Caloric intake (mean) 1286 ± 344      1237 ± 342 Volume ratio at day 7 (%) 80 ± 28      75 ± 30	Diarrhea 7/50 (14) Vomiting 4/50 (8)	Diarrhea 7/51 (14) Vomiting 2/51 (4)
12. Hsu 2009	ICU 18.20 ± 11.80 Hospital 36.0 ± 24.2	ICU 18.20 ± 11.20 Hospital 31.7 ± 21.1	28.5 ± 24.9 (59)	23.8 ± 18.2 (62)	Mean % of daily goal calorie fed 95 ± 5      83 ± 6 Caloric intake (kcal/day) 1658 ± 118      1426 ± 110 Protein (grams/day) 67.9 (4.9)      58.8 (4.9)	Vomiting 1/59 (2) GI bleeding 7/59 (12) Time to reach goal 32.4 (27.1) hrs	Vomiting 8/62 (13) GI bleeding 9/62 (15) Time to reach goal 54.5 (51.4) hrs
13. White 2009	ICU 5.3 (2.73-9.89) 7.12 ± 6.00 (51)	ICU 5.02 (1.98-9.99) 9.10 ± 10.55 (55)	3.93 (2.3-8.38) 5.73 ± 5.29 (51)	3.92 (1.5-8.54) 7.68 ± 9.81 (55)	Caloric intake (median, IQR) 1463 (1232-1804)      1588 (913-1832) Protein intake (median, IQR) 63 (50-78)      69 (45-87)	Time to reach goal 4.1 (3.4-5.0) hrs	Time to reach goal 4.3 (4.0-5.0)
14. Acosta-Escribano 2010	ICU 16 ± 9 (50) Hospital 38 ± 24 (50)	ICU 18 ± 7 (54) Hospital 41 ± 28 (54)	7.3 ± 4 (50)	8.9 ± 4 (54)	Nutritional efficiency (%) 92 ± 7      84 ± 15	High GRVs 3/50 (6) GIT complications 7/50 (14)	High GRVs 15/54 (28) GIT complications 27/54 (47)

15. Davies 2012	<p>ICU 10 (7-15) 12.5 ± 8.6 (91) Hospital 20 (11-33) 28.8 ± 26.1 (91)</p>	<p>ICU 11 (7-16) 12.7 ± 9.8 (89) Hospital 24 (15-32) 27.4 ± 21.1 (89)</p>	<p>8 (6-12) 9.8 ± 6.2 (91)</p>	<p>8 (5-14) 9.7 ± 6.3 (89)</p>	<p>Nutritional efficiency (%) 72 71 p=0.66 Caloric intake (mean) 1497 ± 521 1444 ± 485</p>	<p>Major haemorrhage 2/91 (2) Minor haemorrhage 12/91 (13) Vomiting 30/91 (33) Aspiration 5/91 (5) Diarrhea 26/91 (29) Abdom distention 16/91 (18)</p>	<p>Major haemorrhage 2/89 (2) Minor haemorrhage 3/89 (3) Vomiting 30/89 (30) Aspiration 4/89 (5) Diarrhea 26/89 (30) Abdom distention 18/89 (20)</p>
16. Friedman 2015	<p>ICU 10 (7-21) (54)</p>	<p>ICU 12 (8-20) (61)</p>	<p>4 (2-11) (54)</p>	<p>7 (3-13) (61)</p>	<p>NR</p>	<p>Cost, US\$ 1163 Diarrhea 15/54 (28) Vomiting 14/54 (26) Constipation 9/54 (17)</p>	<p>Cost, US\$ 467 Diarrhea 11/61 (18), p=0.306 Vomiting 18/61, p=0.826 Constipation 14/61 (23), p=0.544</p>
17. Wan 2015	<p>ICU 12.2 ± 0.7 (35)</p>	<p>ICU 17.1 ± 1.0 (35)</p>	<p>5.2 ± 0.3 (35)</p>	<p>8.5 ± 0.5 (35)</p>	<p>NR</p>	<p>Cost 5203 ± 247 Diarrhea 9/35 Reflux 1/35</p>	<p>Cost 7786 ± 555, P &lt;0.01 Diarrhea 9/35 Reflux 14/35, P &lt;0.01</p>

C. Random: concealed randomization  
ITT: intent to treat  
† presumed ICU mortality unless otherwise specified  
‡ refers to the # of patients with infections unless specified

± ( ) : mean ± Standard deviation (number)  
( - ) : median (range)  
NR: not reported



Figure 1. Mortality

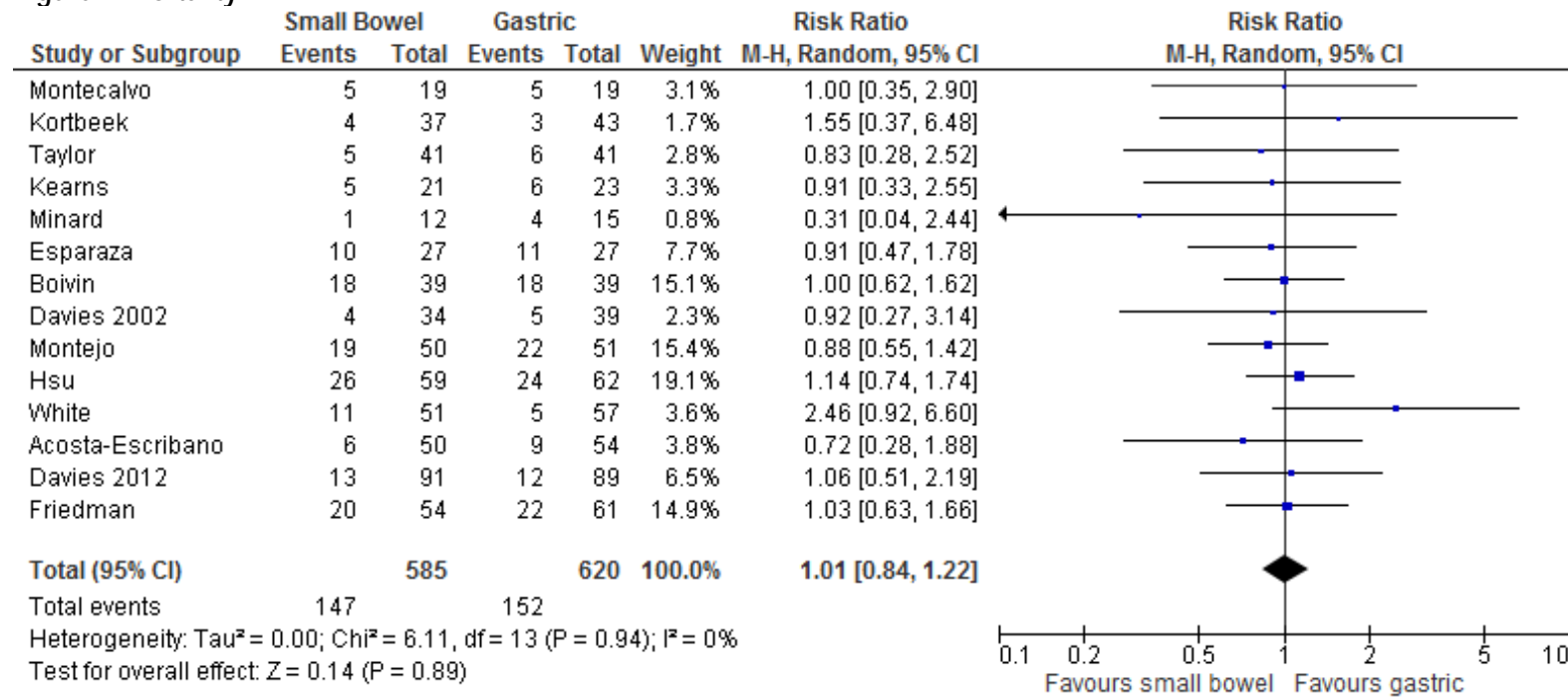


Figure 2. Mortality (excluding Taylor and Minard)

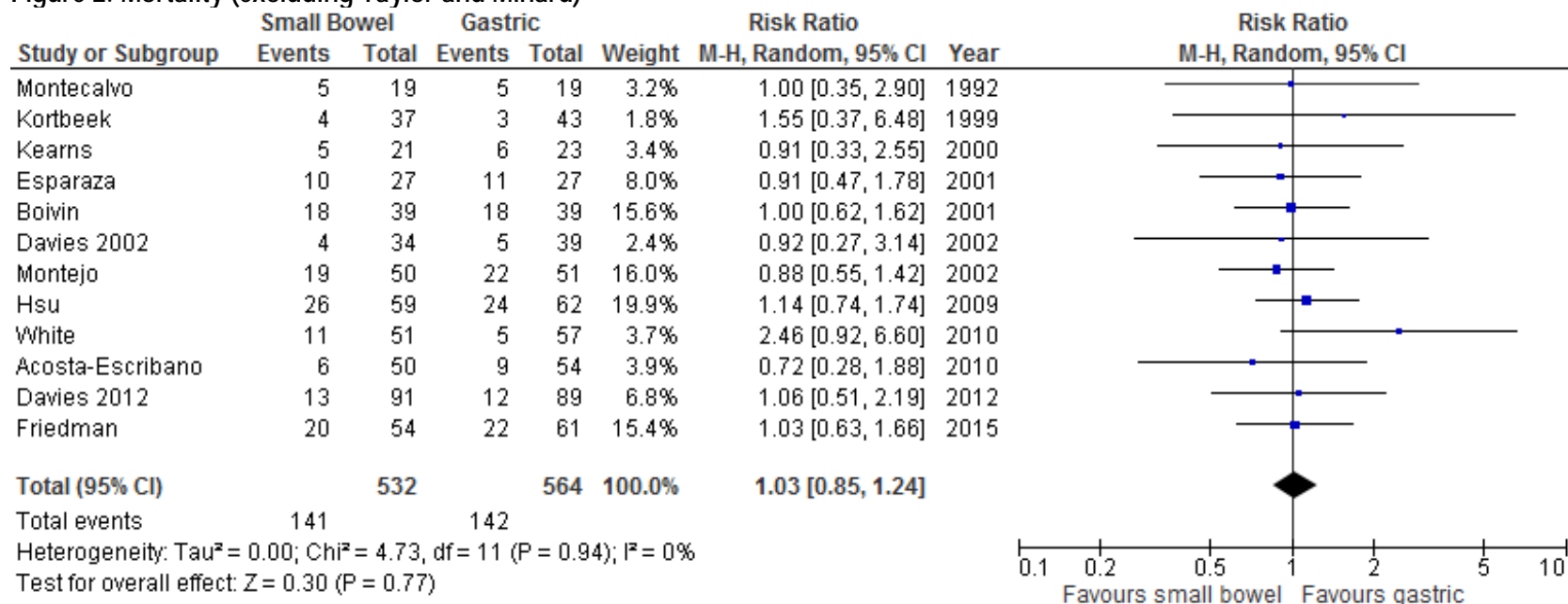


Figure 3. Pneumonia

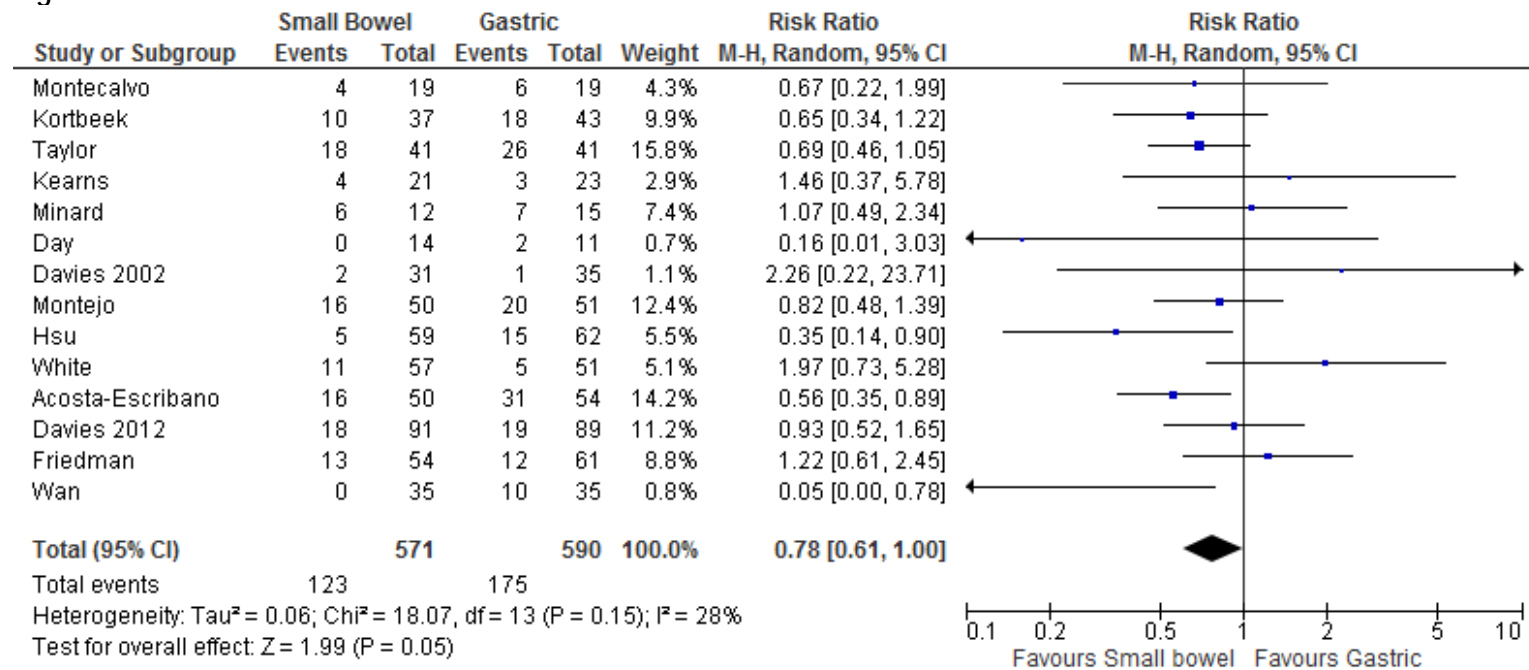


Figure 4. Pneumonia (excluding Taylor and Minard)

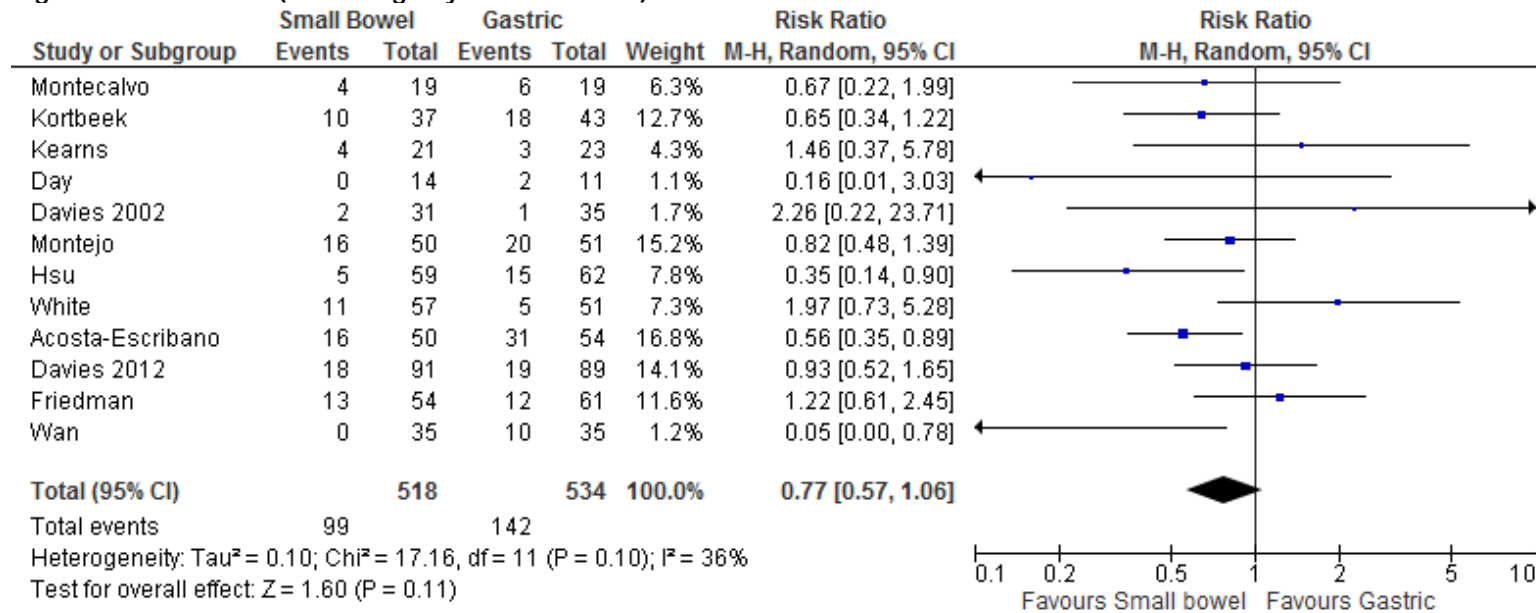


Figure 5. ICU LOS

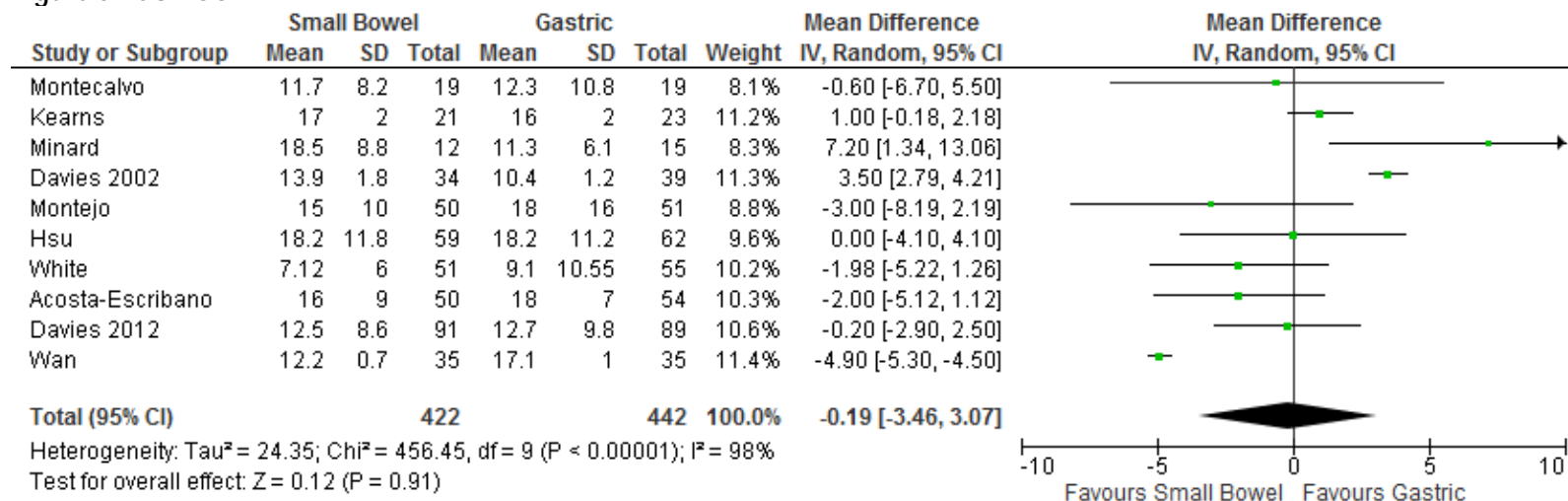


Figure 6. ICU LOS (excluding Minard)

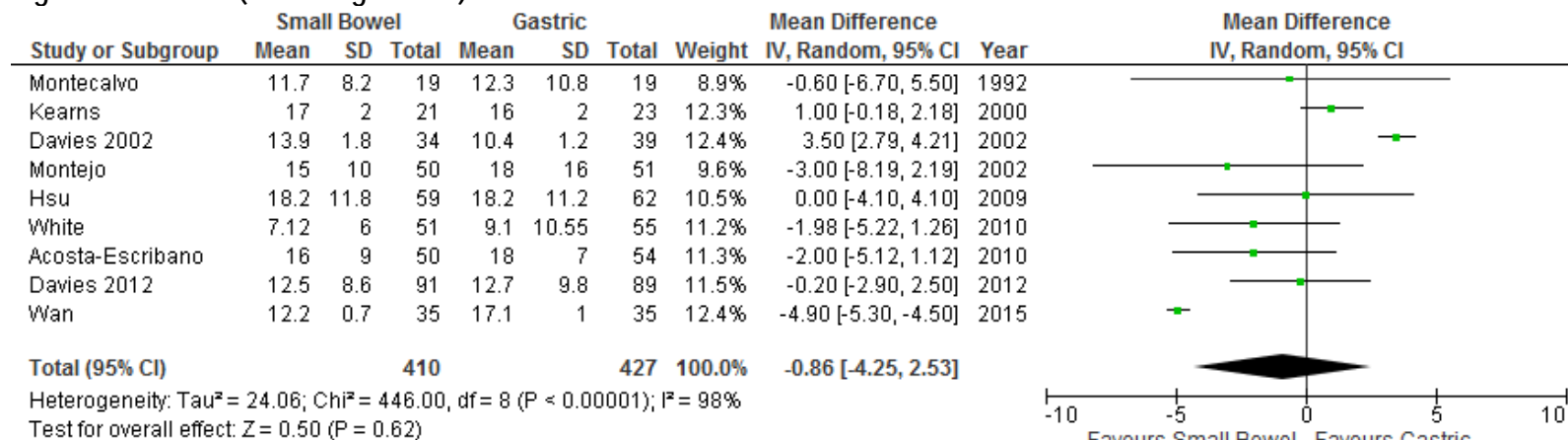


Figure 7. Hospital LOS

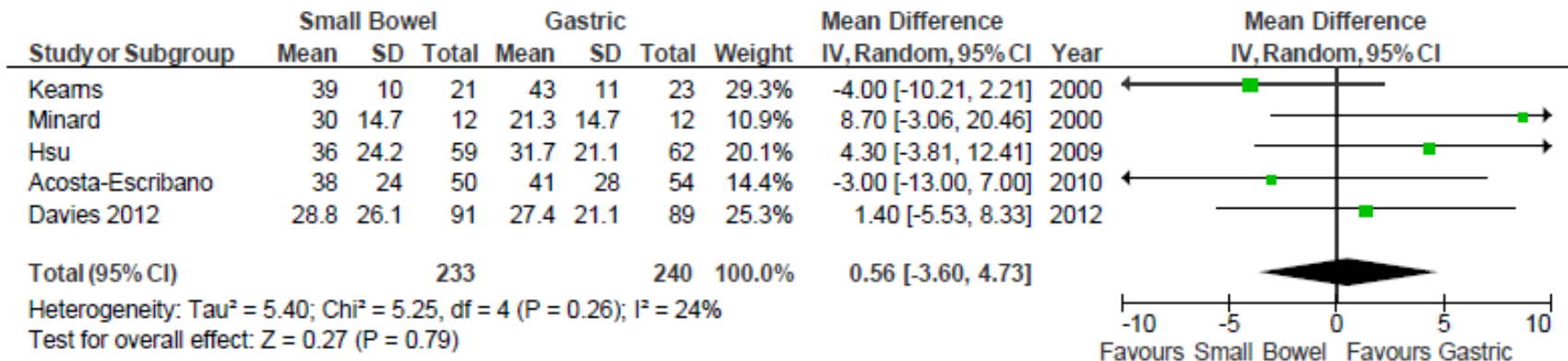


Figure 8. Duration of ventilation

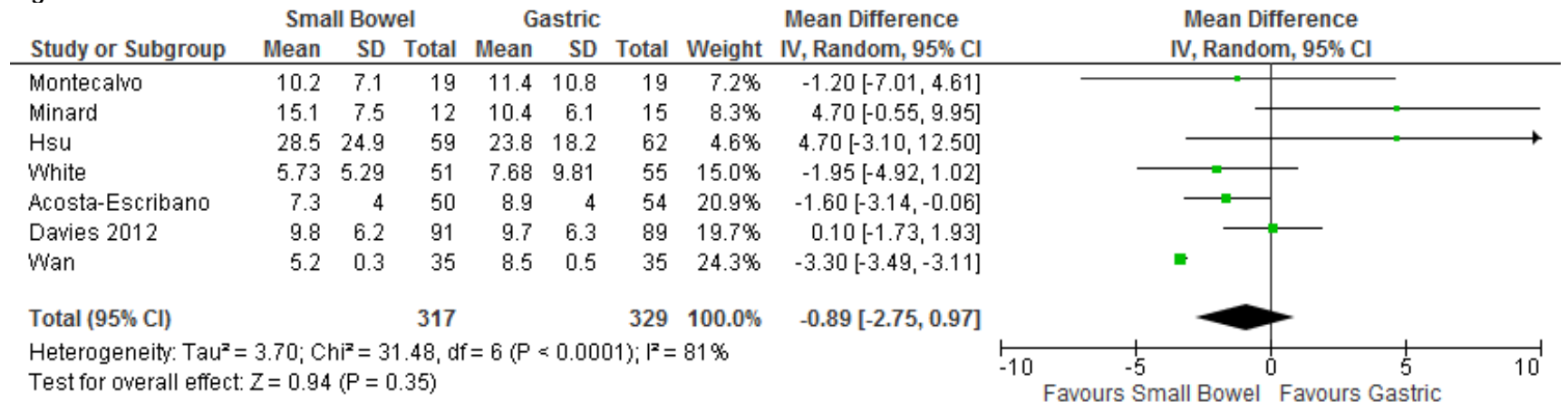


Figure 9. Nutritional efficiency (%)

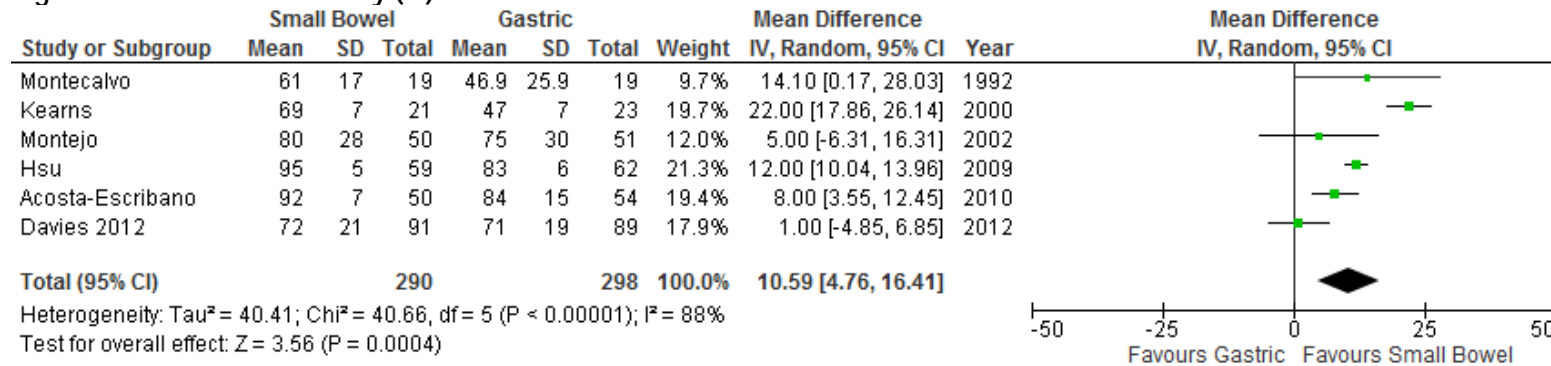


Figure 10. Time to reach EN target

