

CQ5-1 (GRADE)

P: Patients with sepsis/ septic shock/ infection

I: Use of IVIG

C: Placebo or not use of IVIG

O: Mortality, length of ICU stay, serious adverse events

Certainty assessment							№ of patients		Effect		Certainty	Importance
№ of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Treatment	Placebo	Relative (95% CI)	Absolute (95% CI)		
Mortality (All RCT)												
9	randomised trials	serious	not serious	serious	not serious	none	232/769 (30.2%)	271/709 (38.2%)	RR 0.72 (0.58 to 0.90)	107 fewer per 1,000 (from 161 fewer to 38 fewer)	⊕⊕○○ Low	CRITICAL
Mortality (Low RoB)												
3	randomised trials	not serious	not serious	serious	not serious	none	139/381 (36.5%)	131/364 (36.0%)	RR 1.02 (0.84 to 1.23)	7 more per 1,000 (from 58 fewer to 83 more)	⊕⊕⊕○ Moderate	CRITICAL
Length of ICU stay												
4	randomised trials	serious	not serious	serious	not serious	none	233	202	-	MD 1.1 day fewer (from 5.44 fewer to 3.25 more)	⊕⊕○○ Low	CRITICAL
Serious adverse events												
2	randomised trials	not serious	not serious	serious	very serious	none	15/371 (4.0%)	15/353 (4.2%)	RR 0.97 (0.45 to 2.08)	1 fewer per 1,000 (from 23 fewer to 46 more)	⊕○○○ Very low	CRITICAL

	JUDGEMENT						
PROBLEM	No	Probably no	Probably yes	Yes		Varies	Don't know
DESIRABLE EFFECTS	Trivial	Small	Moderate	Large		Varies	Don't know
UNDESIRABLE EFFECTS	Large	Moderate	Small	Trivial		Varies	Don't know
CERTAINTY OF EVIDENCE	Very low	Low	Moderate	High			No included studies
VALUES	Important uncertainty or variability	Possibly important uncertainty or variability	Probably no important uncertainty or variability	No important uncertainty or variability			
BALANCE OF EFFECTS	Favors the comparison	Probably favors the comparison	Does not favor either the intervention or the comparison	Probably favors the intervention	Favors the intervention	Varies	Don't know
ACCEPTABILITY	No	Probably no	Probably yes	Yes		Varies	Don't know
FEASIBILITY	No	Probably no	Probably yes	Yes		Varies	Don't know

CQ5-2-1 (GRADE)

P: Septic patients due to streptococcal infection

I: Use of IVIG

C: Placebo or not use of IVIG

O: Mortality, length of ICU stay, serious adverse events

Certainty assessment							№ of patients		Effect		Certainty	Importance
№ of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Treatment	Placebo	Relative (95% CI)	Absolute (95% CI)		
Mortality (STSS)												
1	randomised trials	not serious	not serious	serious	very serious	none	2/10 (20.0%)	4/11 (36.4%)	RR 0.55 (0.13 to 2.38)	164 fewer per 1,000 (from 316 fewer to 502 more)	⊕○○○ Very low	CRITICAL
3	Observational study	very serious	not serious	serious	serious	none	11/58 (19.0%)	50/115 (43.5%)	RR 0.42 (0.25 to 0.73)	252 fewer per 1,000 (from 326 fewer to 117 fewer)	⊕○○○ Very low	CRITICAL
Mortality (Study which use of CLDM)												
1	randomised trials	not serious	not serious	serious	very serious	none	1/8 (12.5%)	3/10 (30.0%)	RR 0.42 (0.05 to 3.28)	174 fewer per 1,000 (from 285 fewer to 684 more)	⊕○○○ Very low	CRITICAL
4	observational study	serious	not serious	serious	serious	none	13/80 (16.3%)	29/95 (30.5%)	RR 0.53 (0.30 to 0.94)	143 fewer per 1,000 (from 214 fewer to 18 fewer)	⊕○○○ Very low	CRITICAL
Sever adverse events												
1	randomised trials	not serious	not serious	serious	very serious	none	8/50 (16.0%)	11/50 (22.0%)	RR 0.73 (0.32 to 1.65)	59 fewer per 1,000 (from 150 fewer to 143 more)	⊕○○○ Very low	CRITICAL

	JUDGEMENT						
PROBLEM	No	Probably no	Probably yes	Yes		Varies	Don't know
DESIRABLE EFFECTS	Trivial	Small	Moderate	Large		Varies	Don't know
UNDESIRABLE EFFECTS	Large	Moderate	Small	Trivial		Varies	Don't know
CERTAINTY OF EVIDENCE	Very low	Low	Moderate	High			No included studies
VALUES	Important uncertainty or variability	Possibly important uncertainty or variability	Probably no important uncertainty or variability	No important uncertainty or variability			
BALANCE OF EFFECTS	Favors the comparison	Probably favors the comparison	Does not favor either the intervention or the comparison	Probably favors the intervention	Favors the intervention	Varies	Don't know
ACCEPTABILITY	No	Probably no	Probably yes	Yes		Varies	Don't know
FEASIBILITY	No	Probably no	Probably yes	Yes		Varies	Don't know

CQ5-2-2 (UnGRADE)

P: Septic patients due to staphylococcus aureus

I: Use of IVIG

C: Placebo or not use of IVIG

O: Mortality, length of ICU stay, serious adverse events

	JUDGEMENT						
PROBLEM	No	Probably no	Probably yes	Yes		Varies	Don't know
DESIRABLE EFFECTS	Trivial	Small	Moderate	Large		Varies	Don't know
UNDESIRABLE EFFECTS	Large	Moderate	Small	Trivial		Varies	Don't know
CERTAINTY OF EVIDENCE	Very low	Low	Moderate	High			No included studies
VALUES	Important uncertainty or variability	Possibly important uncertainty or variability	Probably no important uncertainty or variability	No important uncertainty or variability			
BALANCE OF EFFECTS	Favors the comparison	Probably favors the comparison	Does not favor either the intervention or the comparison	Probably favors the intervention	Favors the intervention	Varies	Don't know
ACCEPTABILITY	No	Probably no	Probably yes	Yes		Varies	Don't know
FEASIBILITY	No	Probably no	Probably yes	Yes		Varies	Don't know