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Additional File 2: Case Report Form for Methodological Quality of animal research study.

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Title:	
Journal: CCM, ICM, AJRCCM Year; Volume: page numbers:	

Variable	Data	Comments
Title: accurate and concise	Yes or No	If no: randomization; blinding;
		intervention; other.
Abstract: accurate	Yes or No	If no:
		-Omission: background, objectives,
		methods, results, conclusions
		-Inaccuracy: background, objectives,
		methods, results, conclusions.
Introduction:	-	_
Objectives/hypotheses in PICO	Yes or No	-
format		
Number of animals used	-	-
Stated in methods section		-
Any extra in results section	Yes or No	If yes: answer bullets below
-how many more		If number not clear: <5, 5 or more,
		unclear.
Number in the largest group		-
Methods:	-	-
Design:	_	-
Randomized	Yes or No	If no: alternation; dates; picked out of
		cage; historical control; not specified,
		other
		If yes: answer the bullets below
-Procedure described	Yes or No	Table; computer; other
-Block randomization	Yes or No	If yes: size of blocks:
-Stratified randomization	Yes or No	If yes: number of characteristics
-Factorial	Yes or No	If yes: number of factors
-Allocation concealment	Yes or No	If yes: third party; envelopes; other
Blinding	Yes or No	If yes: answer the bullets below.
-disease induction	Yes, No, N/A, or	_
	Unclear	
-treatment/intervention	Yes, No, Some, or	-
	Unclear	
-outcome assessment	All, Some, or	If none or some: any subjective
	None	outcomes not blinded- Yes or No
Experimental Unit	Yes or No	Group; Individual; Part of individual
Eligibility criteria	Yes or No	-
Study procedures:	•	-
Sepsis model	Yes or No	If yes: answer the bullets below.

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-supportive therapy	Yes or No	If yes: fluids; antibiotics; other
-intercurrent illnesses	Yes or No	If yes: describe
-model relevance	-	Focal or bolus; \leq or $>$ 24 hours;
		treatment intervention is pre- or post-
		sepsis induction
Preparation prior to experiment	Yes or No	If yes: acclimation; habituation;
		quarantine
Staff description	Yes or No	If yes: number; training in induction;
-		intervention; monitoring; outcome
Animal description:	-	-
Species	Yes or No	
Strain	Yes or No	-
Sex	Yes or No	
Age	Yes or No	
Developmental stage	Yes or No	If yes: neonate; juvenile; adult
Weight	Yes or No	-
Strain, sex, and either weight or	Yes or No	-
age		
Source	Yes or No	If yes: commercial or local; colony;
		transport; previous use
Genotype	Yes or No	-
Statistics:	-	1 - 2
Sample size calculated	Yes or No	If cluster trial: cluster adjustment YorN
		If sample size calculated: explicit
		mention of alpha, beta/power,
		minimally important difference.
Primary outcome specified	Yes or No	If yes: answer the bullets below.
-number of primary outcomes	-	Number
-statistical method specified	Yes or No	-
Secondary outcomes specified	Yes or No	If yes: answer the bullets below.
-number of secondary outcomes	-	Number: none, <5, 5-10, >10.
-statistical methods specified	Yes or No	-
Sub-groups to be analyzed?	Yes or N/A	If yes: answer the bullets below.
-how many?	-	Number
Results:	-	
baseline characteristics	Yes or No	If yes: answer the bullets below.
compared in table		,
-at least two demographics	Yes or No	
-at least two physiology	Yes or No	
Primary outcome analyzed	Yes or No or N/A	If yes: answer the bullets below.
-numbers with denominators	Yes or No	-
-unaccounted numbers	Yes, No, or	-
	Unclear	
-any excluded from analysis	Yes or No or	If yes: number stated; reasons stated
50 ₹	Unclear	

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numbers in tables/searchs	Yes or No	Ī
-numbers in tables/graphs		If a company of the second of
-all pre-specified reported	Yes or No	If no: number not reported
-measure of precision	Yes or No	If yes: CI; SD; SE
-absolute and relative effect	Yes or No	-
-number positive outcome	-	Number
-number negative or neutral	-	Number
outcome		
-intention to treat analysis	Yes or No	-
-pre-specified subgroups	Yes or No or N/A	If yes: test of interaction done YorN
analyzed		
-post-hoc subgroups analyzed	Yes or N/A	If yes: number; test of
		interaction done YorN
Secondary outcomes analyzed	Yes or No	If yes: answer the bullets below as they
		apply to the majority of the outcomes.
-numbers with denominators for	Yes or No	-
most		
-unaccounted numbers	Yes or No or	-
	Unclear	
-any excluded from analysis	Yes or No or	If yes: number stated; reasons stated
	Unclear	,
-numbers in most tables/graphs	Yes or No	-
-all pre-specified important	Yes or No	-
outcomes reported		
-measure of precision	Yes or No	If yes: CI; SD; SE
-absolute and relative effect	Yes or No	-
-mostly positive outcomes	Yes or No	-
-any negative outcome (obvious,	Yes or No	_
or explicitly stated)		
-number post-hoc outcomes (not	_	Number: none; <5; 5-10; >10
pre-specified in methods)		
-number of statistical	_	<5; 5-20; 21-40; >40
comparisons		2, 2 20, 21 10, 10
-pre-specified subgroups	Yes or No or N/A	If yes: number ; test of
analyzed	105 01 110 01 11/11	interaction done YorN
-post-hoc subgroups analyzed	Yes or N/A	If yes: number ; test of
-post-noe subgroups analyzed	I CS OI IV/A	interaction done YorN.
Any toxicity or lack of toxicity	Yes, No, or N/A	-
mentioned or reported	105, 140, 01 14/11	
Discussion:	_	
Internal validity: limitations	Yes or No	If yes: sample size; bias; multiple
discussed	105 01 110	comparisons.
External validity: to humans	Yes or No	If yes: any limitation to generalizing to
discussed	1 63 01 110	humans mentioned- Yes or No or Vague
Funding source described	Yes or No	If yes: answer the bullets below.
	Yes or No	11 yes. answer the bullets below.
-government source	I es of No	<u> </u>

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-foundation charity	Yes or No	-
-industry company	Yes or No	-
-any role in analysis or reporting	Yes or No or NR	-
Any Comments:		

Notes:

- -circle correct response for Yes or No
 -circle all that apply for sub-questions in last column
 -fill in underlined space as applicable with numbers or names