## Reconciliation of results and studies included

	Current (June, 2018) (Source: First figure online*)	Rygård, May 2018 (Source: Figure 3) PMID <u>29761216</u>	<b>Gibbison, 2017</b> PMID: <u>28351429</u>	Volbeda, 2015 (Source: Figure 2) PMID <u>26100123</u>	Annane (Cochrane), 2015 (Source: Analysis 1.5) PMID 26633262
Outcome(s)					
Efficacy: mortality	Low dose steroids for 7 days or less for septic shock requiring vasopressors: RR 0.90 (95% CI 0.81 to 0.99)	Any dose or duration of steroids: RR 0.98 (95% CI 0.89 to 1.08)	Any dose or duration of steroids: No significant reduction in mortality.	RR 0.89 (95% CI 0.74-1.08)	Low-dose steroids for septic shock: RR 0.88 (95% CI 0.78 to 0.87, fixed-effect model)
Heterogeneity	0% (95% CI 0% to 59%)	O% (CI not provided)	Network meta-analysis	O% (CI not provided)	42% (CI not provided)
Studies included	d				
Cooperative Study Group (CSG), 1963 DOI: <u>link</u>	No (Did not use low-dose corticosteroids)	Yes	Yes	No	Yes
Schumer, 1976 PMID <u>786190</u>	No (Did not use low-dose corticosteroids)	No	Yes	Yes	Yes
Sprung, 1984 PMID <u>6384785</u>	No (Did not use low-dose corticosteroids)	No	Yes	Yes	Yes
Bone, 1987 PMID <u>3306374</u>	No (Did not use low-dose corticosteroids)	Yes	Yes	Yes	Yes
Veterans Administration Systemic Sepsis Cooperative Study Group (VASSCSG), 1987 PMID 2888017	No (Did not use low-dose corticosteroids)	No	Yes	Yes	Yes

Luce, 1988   No					·	
Bollaert, 1999   Yes			No	Yes	Yes	Yes
Bollaert, 1999	PMID <u>3202402</u>	I				
PMID 9559600   Yes   Y		·				
Briegel, 1999	· ·	Yes	Yes	Yes	Yes	Yes
PMID 30321651	PMID <u>9559600</u>					
Chawla, 1999   PMID NA. Data from Cochrane, 2015 Link	Briegel, 1999	Yes	Yes	Yes	Yes	Yes
PMID NA. Data from Cochrane, 2015 Link	PMID <u>10321661</u>					
From Cochrane, 2015   Link	Chawla, 1999	Yes	Yes	Yes	No	Yes
2015 Link	PMID NA. Data					
Annane (Ger-Inf-05), 2002	from Cochrane,					
Inf-05), 2002   PMID 12186604   PMID 12186604   PMID 12186604   PMID 12183187   PMID 12133187   PMID 12133187   PMID 12133187   PMID 12133187   PMID 12133187   PMID 12133187   PMID 15133187   PMID 1513318	2015 <u>Link</u>					
PMID 12186604   No (most patients did not septic shock)   No (most patients did not septic shock)   No (most patients did not septic shock)   No (most patients did not have sepsis or septic shock)   No (most patients did not have sepsis or septic shock)   No (most patients did not have sepsis or septic shock)   No (most patients did not have sepsis or septic shock)   No (most patients did not have sepsis or septic shock)   PMID 15557131	Annane (Ger-	Yes	Yes	Yes	Yes	Yes
Yildz, 2002 PMID 12133187         No (most patients did not septic shock)         No         Yes         No         No           Confalonieri, 2005 PMID 15557131         No (most patients did not have sepsis or septic shock)         No         No         No         Yes         Yes           PMID 16276166         Yes         Yes         Yes         Yes         Yes           PMID 16276166         Yes         Yes         No         No         No         Yes           PMID NA. Data from Cochrane, 2015         No         No         Yes         Yes         Yes           Rinaldi, 2006 PMID 16850006         No         No         No         Yes         Yes         Yes           Cicarelli, 2007 PMID 17992396         Yes         Yes         Yes         Yes         Yes/No (Included in analysis 1.4 but not in 1.5)         No	Inf-05), 2002					
PMID 12133187 septic shock)  Confalonieri, 2005 PMID 15557131  Oppert, 2005 Yes Yes Yes Yes Yes Yes PMID 16275166  Tandam, 2005 PMID NA. Data from Cochrane, 2015  Rinaldi, 2006 PMID 16850006  Cicarelli, 2007 PMID 17992396  Kurugundla, 2008 (Did not use low-dose corticosteroids)  Sprung (CORTICUS),	PMID <u>12186604</u>					
Confalonieri, 2005 PMID 15557131 Oppert, 2005 PMID 16276166 PMID 1627616	Yildz, 2002	No (most patients did not	No	Yes	No	No
Confalonieri, 2005	PMID <u>12133187</u>	septic shock)				
2005   PMID 15557131   have sepsis or septic shock)   PMID 15557131     Oppert, 2005   Yes   Yes   Yes   Yes   Yes   Yes     PMID 16276166   Yes   Yes   Yes   Yes   Yes     PMID 16276166   Yes   Yes   Yes   No   No     PMID NA. Data from Cochrane, 2015     Rinaldi, 2006   PMID 16850006   Shock)   PMID 16850006     Cicarelli, 2007   Yes   Yes   Yes   Yes   Yes   Yes/No (Included in analysis 1.4 but not in 1.5)     Kurugundla, 2008   (Did not use low-dose corticosteroids)   Yes   Yes   Yes   Yes   Yes     Sprung (CORTICUS),   Yes   Ye	Confalonieri	·	No	No	Vec	Voc
PMID 15557131   Sprung (CORTICUS),   PMID 15576156   PMID 15576156   PMID 162576166   PMID 16257666   PMID 16257666   PMID 16257666   PMID 16257666   PMID 1		•	140	NO	163	163
Oppert, 2005 PMID 16276166     Yes     Yes     Yes     Yes       Tandam, 2005 PMID NA. Data from Cochrane, 2015     Yes     Yes     No     No     Yes       Rinaldi, 2006 PMID 16850006     No (patients did not have septic shock)     No     Yes     Yes     Yes       Cicarelli, 2007 PMID 17992396     Yes     Yes     Yes     Yes/No (Included in analysis 1.4 but not in 1.5)       Kurugundla, 2008     No (Did not use low-dose corticosteroids)     Yes     Yes     Yes     Yes       Sprung (CORTICUS),     Yes     Yes     Yes     Yes		lave sepsis of septic shock)				
PMID 16276166       Yes       Yes       No       No       Yes         PMID NA. Data from Cochrane, 2015       No       No       Yes       Yes         Rinaldi, 2006       No (patients did not have septic shock)       No       Yes       Yes       Yes         Cicarelli, 2007       PMID 17992396       Yes       Yes       Yes       Yes/No (Included in analysis 1.4 but not in 1.5)         Kurugundla, 2008       No (Did not use low-dose corticosteroids)       Yes       No       No       No         Sprung (CORTICUS),       Yes       Yes       Yes       Yes       Yes		Yes	Yes	Yes	Yes	Yes
Tandam, 2005 PMID NA. Data from Cochrane, 2015 Rinaldi, 2006 PMID 16850006 Cicarelli, 2007 PMID 17992396 Kurugundla, 2008 CiDid not use low-dose corticosteroids)  Yes  Yes  No No Yes		163	103	103	163	163
PMID NA. Data from Cochrane, 2015  Rinaldi, 2006 PMID 16850006  Cicarelli, 2007 PMID 17992396  Kurugundla, 2008  Sprung (CORTICUS),  Rinaldi, 2006 No (patients did not have septic shock)  Yes  Yes  Yes  Yes  Yes  Yes  Yes  Ye		Ves	Ves	No	No	Ves
from Cochrane, 2015NoNoYesYesYesRinaldi, 2006 PMID 16850006 PMID 16850006NoNoYesYesYesCicarelli, 2007 PMID 17992396YesYesYesYesYesKurugundla, 2008 2008NoYesNoNoNoSprung (CORTICUS),YesYesYesYesYes		163	103	110	140	1.63
2015Rinaldi, 2006 PMID 16850006No (patients did not have septic shock)No YesYesYesYesCicarelli, 2007 PMID 17992396YesYesYesYesYes/No (Included in analysis 1.4 but not in 1.5)Kurugundla, 2008No (Did not use low-dose corticosteroids)YesNoNoNoSprung (CORTICUS),YesYesYesYesYes						
Rinaldi, 2006 PMID 16850006 (patients did not have septic shock)  Cicarelli, 2007 PMID 17992396  Kurugundla, 2008  (Did not use low-dose corticosteroids)  No Yes						
PMID 16850006 (patients did not have septic shock)  Cicarelli, 2007 PMID 17992396 PMID 17992396  Kurugundla, No (Did not use low-dose corticosteroids)  Sprung (CORTICUS),  (patients did not have septic shock)  Yes  Yes  Yes  Yes  Yes  Yes  No  No  No  No  No  Yes  Yes  Yes  Yes  Yes  Yes  Yes  Ye		No	No	Yes	Yes	Yes
Cicarelli, 2007 PMID 17992396  Kurugundla, 2008  Cicarelli, 2007 PMID 17992396  Cicarelli, 2007 PMID 17992396  Sprung (CORTICUS),  Keres  Yes  Yes  Yes  Yes  Yes  Yes  Yes	· ·			. 55	. 55	. 55
Cicarelli, 2007 PMID 17992396  Yes Yes Yes Yes Yes Yes/No (Included in analysis 1.4 but not in 1.5)  Kurugundla, 2008 (Did not use low-dose corticosteroids)  Yes Yes Yes Yes Yes Yes Yes Yes Yes Ye						
PMID 17992396 Kurugundla, 2008 (Did not use low-dose corticosteroids)  Yes Yes Yes Yes Yes Yes Yes Yes Yes Ye	Ciacrelli 2007	·	Ve-	V	V	Vec/NI-
Kurugundla, 2008 (Did not use low-dose corticosteroids)  Yes Yes Yes Yes Yes Yes Yes Yes Yes Ye		Yes	Yes	Yes	Yes	-
Kurugundla, 2008 (Did not use low-dose corticosteroids)  Sprung (CORTICUS),	PIVIID <u>17992396</u>					
2008 (Did not use low-dose corticosteroids)  Sprung (CORTICUS),  Yes Yes Yes Yes Yes						-
corticosteroids)  Sprung Yes Yes Yes Yes Yes Yes Yes	_		Yes	No	No	No
Sprung Yes Yes Yes Yes Yes	2008					
(CORTICUS),		corticosteroids)				
(CORTICUS),	Sprung	Yes	Yes	Yes	Yes	Yes

PMID <u>18184957</u>					
Hu, 2009 PMID 19751560. Data from Cochrane, 2015	Yes	Yes	Yes	Yes	Yes
Meduri, 2009 PMID <u>17426195</u>	No (most patients did not septic shock)	Yes	Yes	Yes	Yes
Snijders, 2010 PMID 20133929	No (most patients did not septic shock)	No	Yes	No (see footnote in their Table 2)	Yes
Arabi, 2010 PMID <u>21059778</u>	Yes	Yes	Yes	Yes	Yes
Meijvis. 2011 PMID 21636122	No (most patients did not septic shock)	No	Yes	No	Yes
Sabry, 2011	No (most patients did not septic shock)	No	Yes	No	Yes
Yildz, 2011 PMID <u>22973341</u>	No (most patients did not septic shock)	No	Yes	No	No
Liu, 2012 PMID <u>23158856</u>	No (most patients did not septic shock)	No	Yes	No	Yes
Gordon, 2014 PMID <u>24557425</u>	Yes	Yes	Yes	Yes	Yes
Mirea, 2014 <u>PMC4068893</u>	Yes †	Yes	No	No	No
Rezk, 2014 PMID NA	No (most patients did not septic shock)	No	No	No	NO
Torres, 2014	No (most patients did not septic shock)	Yes	No	No	Yes
Gordon (VANISH), 2016 PMID <u>27483065</u>	Yes †	Yes	No	NA	NA
Tongyoo, 2016 PMID <u>27741949</u>	Yes	Yes	No	NA	NA

Key (HYPRESS),	No	No	No	NA	NA
2016	(patients did not have septic				
PMID <u>27695824</u>	shock)				
Lv, 2017	Yes	Yes	No	NA	NA
PMID <u>28615145</u>					
Ventatesh	Yes	Yes	NA	NA	NA
(ADRENAL),					
2018					
PMID <u>29347874</u>					
Annane	Yes	Yes	NA	NA	NA
(APROCCHSS),					
2018					
PMID <u>29490185</u>					
	Current	Rygård, 2017	Gibbison, 2017	Volbeda, 2015	Annane (Cochrane),
	(Source: First figure online*)	(Source: Figure 3)	PMID: <u>28351429</u>	(Source: Figure 2)	2015
		PMID <u>29761216</u>		PMID <u>26100123</u>	(Source: Analysis 1.5)
					PMID <u>26633262</u>

## Notes:

<sup>\* &</sup>lt;a href="http://openmetaanalysis.github.io/sepsis-steroids">http://openmetaanalysis.github.io/sepsis-steroids</a>

<sup>†</sup> The current review, when published 06/2018, did not identify the VANISH study in title screening and was not aware of the mortality data obtained by the Rygård review and published in 05/2021 between closing and publishing our review in 06/2018. These two trials and this footnote were added 07/2021.