

	Butcher Meshke & Sheppard 2004	Eisner et al 2008	Smith et al 2010	Roig et al 2010	Rocco et al 2011	Roig et al 2011	Ozalevli et al 2011	Beauchamp et al 2012	Janssens et al 2013	Janssens et al 2014	Mkacher et al 2014	Pedrozo & Silveira 2015	Yumrutepe et al 2015	Crisan et al 2015	Oliveira et al 2015	Tudorache et al 2015	Smith, Chang & Hodges 2016	Voica et al 2016	Iwakura et al 2016	De Castro et al 2016	Porto et al 2017	Oliveira et al 2017	Morlino et al 2017
1. Hypothesis /aim/ objectives clearly described?	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes
2. Main outcomes clearly described in introduction/ methods?	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes
3. Characteristics of patients included clearly described?	no	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes
4. Interventions of interest clearly described?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a
5. Distributions of confounders in each group to be compared clearly described?	yes	yes	yes	UD NR	yes	UD NR	yes	yes	yes	yes	yes	UD NR	yes	UD NR	yes	yes	yes	UD NR	yes	UD NR	yes	yes	yes
6. Main findings clearly described?	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes
7. Estimates of the random variability in the data for the main outcomes provided?	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes

8. All adverse events potentially from the intervention reported?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a
9. Characteristics of patients lost to follow-up described?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a
10. Actual probability values for the main outcomes reported?	yes	yes	yes	yes	no	yes	yes	yes	yes	yes	no	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes
Reporting subscore (/11) (/8)	7	7	8	7	6	7	8	8	8	8	7	7	8	7	8	8	8	8	8	7	7	7	7
11. Subjects asked to participate representative of the entire population?	UD NR	yes	UD NR	yes	UD NR	yes	no	UD NR	UD NR	UD NR	UD NR	yes	UD NR	UD NR	yes	UD NR	UD NR	UD NR	no	UD NR	UD NR	yes	UD NR
12. Subjects who accepted to participate representative of the entire population?	UD NR	no	UD NR	yes	UD NR	no	no	UD NR	UD NR	UD NR	UD NR	no	UD NR	UD NR	UD NR	UD NR	UD NR	UD NR	no	UD NR	UD NR	yes	UD NR
13. Staff, places, facilities where patients were treated representative of the treatment most patients receive?	yes	yes	UD NR	yes	yes	no	yes	UD NR	UD NR	UD NR	UD NR	yes	yes	UD NR	yes	no	UD NR	no	yes	UD NR	yes	yes	UD NR
External validity subscore (/3) (/3)	1	3	0	3	1	3	3	0	0	0	0	2	1	0	2	0	0	1	2	0	1	3	0
14. Attempt made to blind study subjects?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a
15. Attempt made to blind those measuring the main outcomes?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a

16. Any results based on "data dredging" made clear?	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	no	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes
17. Analyses adjust for different lengths of follow-up or follow-up length the same for cases and controls?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a
18. Statistical tests for main outcomes appropriate?	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes
19. Compliance with intervention reliable?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a
20. Main outcome measures used valid and reliable?	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes	yes
Internal validity – bias (/7) (/3)	3	3	3	3	3	3	3	3	3	2	3	3	3	3	3	2	3	3	3	3	3	3	3
21. Subjects from both groups recruited from the same population?	yes	yes	UD NR	yes	UD NR	yes	UD NR	no	UD NR	no	UD NR	UD NR	UD NR	no	yes	UD NR	UD NR	UD NR	yes	no	UD NR	yes	UD NR
22. Subjects from both groups recruited over the same time period?	no	no	UD NR	yes	UD NR	yes	UD NR	UD NR	UD NR	UD NR	UD NR	yes	yes	UD NR	yes	UD NR	UD NR	UD NR	UD NR	UD NR	UD NR	no	UD NR
23. Study subjects randomised to intervention group?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a
24. Randomisation concealed from patients and staff until recruitment complete and irrevocable?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a

25. Adequate adjustment for confounding in the analyses for main findings?	yes	yes	no	yes	no	yes	no	yes	yes	yes	yes	no	no	no	yes	no	no	yes	yes	no	yes	yes	no
26. Losses of patient to follow-up taken into account?	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a	n/a
Internal validity – confounding (selection bias) (/6) (/3)	1	3	1	3	1	1	3	1	1	1	1	1	2	1	3	0	0	1	2	1	1	3	1
Study adequately powered?	no	yes	no	yes	yes	no	no	yes	no	no	yes	yes	no	no	yes	no	no	no	yes	yes	no	yes	no
Power (/1)	0	1	0	0	1	0	0	1	0	0	0	0	0	2	1	0	0	0	1	1	0	1	0
TOTAL SCORE (/28) (/18)	11	14	11	13	11	13	14	12	11	10	10	12	12	12	14	10	11	12	14	11	11	14	10
UD = Unable to determine; NR=not reported; PR=partially reported; n/a= non-applicable																							