Multimedia Appendix III: Risk of Bias Assessment Questions

Taken from the Agency for Health Research and Quality's 2014 Methods Guide for Effectiveness and Comparative Effectiveness Reviews [18]

Randomized Controlled Trials

Risk of Bias	Criterion
Selection Bias	Was the allocation sequence generated adequately (e.g., random number table, computer-generated randomization)?
	Was the allocation of treatment adequately concealed (e.g., pharmacy- controlled randomization or use of sequentially numbered sealed envelopes)?
	Were participants analyzed within the groups they were originally assigned to?
	Does the design or analysis control account for important confounding and
	modifying variables through matching, stratification, multivariable analysis, or
	other approaches?
Performance Bias	Did researchers rule out any impact from a concurrent intervention or an
	unintended exposure that might bias results?
	Did the study maintain fidelity to the intervention protocol?
Attrition Bias	If attrition (overall or differential nonresponse, dropout, loss to follow-up, or
	exclusion of participants) was a concern, were missing data handled
	appropriately (e.g., intention-to-treat analysis and imputation)?
Detection Bias	In prospective studies, was the length of follow-up different between the groups,
	or in case-control studies, was the time period between the intervention/exposure and outcome the same for cases and controls?
	Were the outcome assessors blinded to the intervention or exposure status of
	participants
	Were interventions/exposures assessed/defined using valid and reliable measures,
	implemented consistently across all study participants?
	Were outcomes assessed/defined using valid and reliable measures, implemented
	consistently across all study participants?
Reporting Bias	Were the potential outcomes pre-specified by the researchers? Are all pre- specified outcomes reported?

Cohort and Non-Randomized Controlled Trials

Risk of Bias	Criterion
Selection Bias	Were participants analyzed within the groups they were originally assigned to? Did the study apply inclusion/exclusion criteria uniformly to all comparison groups?
	Did the strategy for recruiting participants into the study differ across study groups?
	Does the design or analysis control account for important confounding and modifying variables through matching, stratification, multivariable analysis, or other approaches?
Performance Bias	Did researchers rule out any impact from a concurrent intervention or an unintended exposure that might bias results?

	Did the study maintain fidelity to the intervention protocol?
Attrition Bias	If attrition (overall or differential nonresponse, dropout, loss to follow-up, or
	exclusion of participants) was a concern, were missing data handled
	appropriately (e.g., intention-to-treat analysis and imputation)?
Detection Bias	In prospective studies, was the length of follow-up different between the groups,
	or in case-control studies, was the time period between the intervention/exposure
	and outcome the same for cases and controls?
	Were the outcome assessors blinded to the intervention or exposure status of
	participants
	Were interventions/exposures assessed/defined using valid and reliable measures,
	implemented consistently across all study participants?
	Were outcomes assessed/defined using valid and reliable measures, implemented
	consistently across all study participants?
	Were confounding variables assessed using valid and reliable measures,
	implemented consistently across all study participants?
Reporting Bias	Were the potential outcomes pre-specified by the researchers? Are all pre-
	specified outcomes reported?