Risk of bias assessment for included studies using Cochrane Collaboration's Tool.

1. Risk of bias assessment for MYRROR (2014)¹

Bias	Authors' judgement	Support for judgement
Random sequence	Unclear risk	Quote: "MYRROR was an
generation (selection bias)		international, phase III,
		multicenter, randomized,
		double-masked, sham-controlled
		study.
		Eligible patients were randomized
		in a 3:1 ratio to receive intravitreal
		aflibercept or sham control
		(stratified by country). "
		The trial was described as
		randomised, but the method of
		sequence generation was not
		specified, we assessed as "
		Unclear risk " .
Allocation concealment	Unclear risk	Not reported
(selection bias)		
Blinding of participants	Low risk	Quote: "MYRROR was an

and personnel		international, phase III,
(performance bias) All		multicenter, randomized,
outcomes		double-masked, sham-controlled
		study."
Blinding of outcome	Low risk	Quote: "MYRROR was an
assessment (detection		international, phase III,
bias)		multicenter, randomized,
All outcomes		double-masked, sham-controlled
		study."
Incomplete outcome data	Low risk	Quote: "In total, 122 patients were
(attrition bias) All		randomized, of whom 91 received
outcomes		intravitreal aflibercept 2.0 mg and
		31 received sham; 122 patients
		were included in the safety set. In
		the full analysis set, 121 patients
		were included (90 patients
		received intravitreal aflibercept 2.0
		mg and 31 received sham). "
		Quote: "According to participant
		flow data on ClinicalTrials.gov, 5
		participants were withdrawn from

			the study and 1 participant did not
			complete visits to week 48 due to
			adverse events, both in the
			aflibercept group. However, only 1
			participant failed to fulfil
			requirements of full analysis set
			after randomisation. "
Selective	reporting	Low risk	All prespecified outcomes were
(reporting bias)			reported.
Other bias		Low risk	No other bias identified.

Ikuno Y, Ohno-Matsui K, Wong TY, et al. Intravitreal Aflibercept Injection in Patients with Myopic Choroidal Neovascularization: The MYRROR Study. Ophthalmology 2015; 122:1220-7.

2. Risk of bias assessment for Parodi et al (2010)²

Bias	Authors'	Support for judgement
	judgement	
Random sequence	Low risk	Quote: " Each patient was randomly
generation (selection bias)		allocated to 1 of the 3 treatment
		groups through a
		computer-generated number. "
Allocation concealment	Unclear risk	Not reported

(selection bias)		
Blinding of participants	Unclear risk	Not reported
and personnel		
(performance bias) All		
outcomes		
Blinding of outcome	Low risk	Quote: "At each scheduled
assessment (detection		examination, a complete
bias)		ophthalmological assessment was
All outcomes		carried out by an investigator who
		had had no previous contact with
		the subject and was unaware of the
		treatment previously
		administered. "
Incomplete outcome data	Low risk	Quote: "Fifty-four patients affected
(attrition bias) All		by juxtafoveal CNV in pathologic
outcomes		myopia were recruited; 4 patients
		were excluded because they could
		not attend the scheduled
		examinations; 3 patients were not
		recruited because they were affected
		by media opacity. "
Selective reporting	Low risk	All prespecified outcomes were

(reporting bias)		reported.
Other bias	Low risk	No other bias identified.

2. Parodi MB, Iacono P, Papayannis A, et al. Laser photocoagulation, photodynamic therapy, and intravitreal bevacizumab for the treatment of juxtafoveal choroidal neovascularization secondary to pathologic myopia. Arch Ophthalmol 2010; 128:437-42.

3. Risk of bias assessment for Moreno et al (2013)^{3,4}

Bias	Authors' judgement	Support for judgement
Random sequence	Low risk	Quote: "The randomisation was
generation (selection bias)		done by the promotor and was
		provided by the IOBA."
		Quote: "We performed a
		multicenter prospective study on
		55 highly myopic eyes from 55
		patients with CNV who were
		randomized to PDT (Group 1) or
		intravitreal bevacizumab (IVB)
		(Group 2)."
Allocation concealment	Low risk	Quote: "The randomisation was
(selection bias)		done by the promotor and was
		provided by the IOBA."

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Blinding of participants	Low risk	Quote: "The study was doubled
and personnel		masked: (the follow-up physician
(performance bias) All		and the optometrist) and the
outcomes		patient were masked."
Blinding of outcome	Low risk	Quote: "The study was doubled
assessment (detection		masked: (the follow-up physician
bias)		and the optometrist) and the
All outcomes		patient were masked."
Incomplete outcome data	High risk	Quote: "Twenty-four eyes in group
(attrition bias) All		1 (86%) and 25 eyes in group 2
outcomes		(92.6%) completed 1 year of
		follow-up and 20 eyes in group 1
		(71.4%) and 22 eyes in group 2
		(78.6%) completed 2 years of
		follow-up."
		The loss to follow-up was > 20%
		at 2 years and no reason was
		reported.
Selective reporting	Low risk	All prespecified outcomes were
(reporting bias)		reported.
Other bias	Low risk	No other bias identified.

^{3.} Ruiz-Moreno JM, López-Gálvez MI, Montero Moreno JA, et al. Intravitreal bevacizumab in

myopic neovascular membranes: 24-month results. Ophthalmology 2013; 120:1510-1.e1.

4. Zhu Y, Zhang T, Xu G, et al. Anti-vascular endothelial growth factor for choroidal neovascularisation in people with pathological myopia. Cochrane Database Syst Rev 2016; 12:CD011160.

4. Risk of bias assessment for RADIANCE (2014)⁵

Bias	Authors' judgement	Support for judgement
Random sequence	Low risk	Quote: "A randomization list was
generation (selection bias)		produced by Novartis Drug Supply
		Management using a validated
		system that automates the random
		assignment of treatment groups to
		randomization numbers in the
		specified ratio."
Allocation concealment	Low risk	Quote: "At enrollment, patients
(selection bias)		received the lowest available
		randomization number that then
		assigned them in a 2:2:1 ratio to 1
		of the 3 treatment groups."
Blinding of participants	Low risk	Quote: "Due to the different
and personnel		appearances and routes of

(performance bias) All		administration between the 2
outcomes		treatments, all patients received
		either sham injection or PDT sham
		in conjunction with the study
		treatment. The PDT sham
		consisted of intravenous injection
		of 5% dextrose solution followed
		by light application of PDT. "
		Quote: "The treating investigator
		was unmasked and administered
		the randomized study medication
		per the protocol; however, they
		were not involved in any other
		aspects of the study and could not
		communicate details of the
		treatment."
Blinding of outcome	Low risk	Quote: "To ensure masking, 2
assessment (detection		investigators were involved at
bias)		each study center. All study
All outcomes		assessments were made by the
		evaluating investigator, VA
		assessor, or other site personnel

		who were masked to the treatment
		assignment. "
Incomplete outcome data	Low risk	Quote: "6(5.7%) patients
(attrition bias) All		discontinued from the study:
outcomes		1(0.9%) unsatisfactory therapeutic
		effect; 1(0.9%) subject withdrew
		consent; 3(2.8%) lost to follow-up;
		1(0.9%) protocol deviation.
		4(3.4%) patients discontinued
		from the study: 2(1.7%) subject
		withdrew consent; 1(0.9%) lost to
		follow-up; 1(0.9%) protocol
		deviation. "
Selective reporting	Low risk	All prespecified outcomes were
(reporting bias)		reported.
Other bias	Low risk	No other bias identified.

5. Wolf S, Balciuniene VJ, Laganovska G, et al. RADIANCE: a randomized controlled study of ranibizumab in patients with choroidal neovascularization secondary to pathologic myopia.

Ophthalmology 2014; 121:682-92.e2.

5. Risk of bias assessment for BRILLIANCE (2019)⁶

Bias	Authors' judgement	Support for judgement
Random sequence	Low risk	Quote: "Eligible patients were
generation (selection bias)		randomized 2:2:1 to one of three
		treatment arms using an interactive
		response technology system (see
		Figure, Supplemental Digital
		Content 3,
		http://links.lww.com/IAE/A901,
		which shows treatment schedule
		and study design)."
Allocation concealment	Low risk	Quote: "Eligible patients were
(selection bias)		randomized 2:2:1 to one of three
		treatment arms using an interactive
		response technology system (see
		Figure, Supplemental Digital
		Content 3,
		http://links.lww.com/IAE/A901,
		which shows treatment schedule
		and study design). "
Blinding of participants	Low risk	Quote: "BRILLIANCE was a
and personnel		12-month, Phase III, randomized,
(performance bias) All		double-masked, multicenter,

outcomes		active-controlled clinical trial."			
		Quote: "For masking purpose,			
		sham ranibizumab or sham vPDT			
		was applied."			
		Quote: "All patients were masked			
		to the study treatment."			
Blinding of outcome	Low risk	Quote: "In addition, to fulfill the			
assessment (detection		masking, there were at least two			
bias)		investigators involved into the			
All outcomes		study: masked (assessing)			
		investigator performing all			
		assessments and capturing data;			
		and an unmasked (treating)			
		investigator administering the			
		randomized study treatment when			
		needed according to the protocol."			
Incomplete outcome data	Low risk	Quote: "9(4.9%) patients			
(attrition bias) All		discontinued from the study in			
outcomes		group 1: 1(0.5%) adverse event;			
		7(3.8%) subject withdrew consent;			
		1(0.5%) lost to follow-up."			
		Quote: "9(4.9%) patients			

		discontinued from the study in				
		group 2: 2(1.1%) adverse event;				
		3(1.6%) subject withdrew consent;				
		2(1.1%) administrative problems;				
		2(1.1%) physician's decision."				
		Quote: "8(8.8%) patients				
		discontinued from the study in				
		group 3: 7(7.7%) subject withdrew				
		consent; 1(1.1%) physician's				
		decision."				
Selective reporting	Low risk	All prespecified outcomes were				
(reporting bias)		reported.				
Other bias	Low risk	No other bias identified.				

6. Chen Y, Sharma T, Li X, et al. Ranibizumab versus verteporfin photodynamic therapy in Asian patients with myopic choroidal neovascularization: BRILLIANCE, a 12-month, randomized, double-masked study. Retina 2019; 39:1985-1994.

6. Risk of bias assessment for Saviano et al (2013)⁷

Bias	Authors' judgement	Support for judgement			
Random sequence	Unclear risk	Quote: "Thirty-four patients we			
generation (selection bias)		included in the study and then			
		randomized into two different			

		treatment groups."				
		The trial was described as				
		randomised, but the method of				
		sequence generation was not				
		specified, we assessed as "				
		Unclear risk " .				
Allocation concealment	Unclear risk	Not reported				
(selection bias)	131					
	I Implementals	Not reported				
Blinding of participants	Unclear risk	Not reported				
and personnel						
(performance bias) All						
outcomes						
Blinding of outcome	Unclear risk	Not reported				
assessment (detection						
bias)						
All outcomes						
Incomplete outcome data	Low risk	No loss to follow-up.				
(attrition bias) All						
outcomes						
Selective reporting	Low risk	All prespecified outcomes were				
(reporting bias)		reported.				
Other bias	Low risk	No other bias identified.				

7. Saviano S, Piermarocchi R, Leon PE, et al. Combined therapy with bevacizumab and photodynamic therapy for myopic choroidal neovascularization: A one-year follow-up controlled study. Int J Ophthalmol 2014; 7:335-9.

7. Risk of bias assessment for Rinaldi et al (2016)⁸

Bias	Authors' judgement	Support for judgement			
Random sequence	Low risk	Quote: "Randomization was			
generation (selection bias)		performed using			
		computer-generated random			
		numbers: each number			
		corresponded to a type of			
		treatment."			
Allocation concealment	Low risk	Quote: "Randomization was			
(selection bias)		performed using			
		computer-generated random			
		numbers: each number			
		corresponded to a type of			
		treatment."			
Blinding of participants	High risk	Quote: "The study was a			
and personnel		prospective, comparative,			
(performance bias) All		interventional, randomized,			
outcomes		openlabel clinical trial."			

Blinding of outcome	High risk	Quote: "The study was a				
assessment (detection		prospective, comparative,				
bias)		interventional, randomized				
All outcomes		openlabel clinical trial."				
Incomplete outcome data	Low risk	Quote: "All patients completed the				
(attrition bias) All		follow-up at 48 weeks."				
outcomes						
Selective reporting	Low risk	All prespecified outcomes were				
(reporting bias)		reported.				
Other bias	Low risk	No other bias identified.				

8. Rinaldi M, Semeraro F, Chiosi F, et al. Reduced-fluence verteporfin photodynamic therapy plus ranibizumab for choroidal neovascularization in pathologic myopia. Graefes Arch Clin Exp Ophthalmol 2017; 255:529-539.

2. Risk of bias summary for included RCTs.

	Random sequence generation (selection bias)	Allocation concealment (selection bias)	Blinding of participants and personnel (performance bias)	Blinding of outcome assessment (detection bias)	Incomplete outcome data (attrition bias)	Selective reporting (reporting bias)	Other bias
BRILLIANCE 2019	•	•	•	•	•	•	•
Moreno 2013	•	•	•	•		•	•
MYRROR 2014	?	?	•	•	•	•	•
Parodi 2010	•	?	?	•	•	•	•
RADANCE 2014	•	•	•	•	•	•	•
Rinaldi 2016	•	•	•		•	•	•
Saviano 2013	?	?	?	?	•	•	•