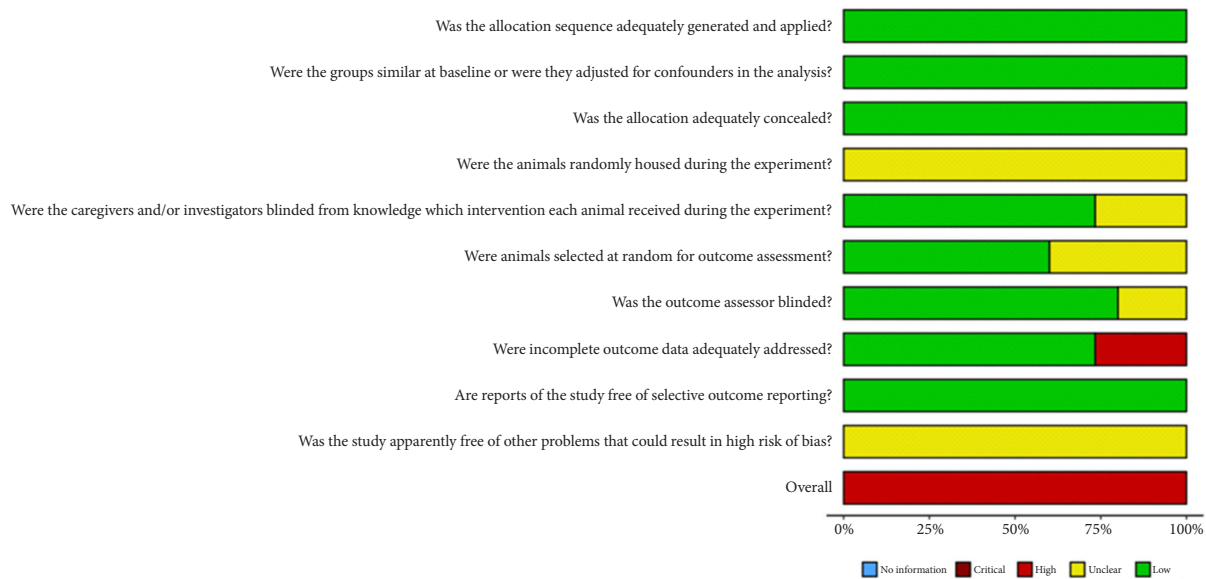


		Risk of bias										
		D1	D2	D3	D4	D5	D6	D7	D8	D9	D10	Overall
Study	Brown, 2011	+	+	+	-	-	-	-	+	+	-	⊗
	Cheng Yu, 2022	+	+	+	-	+	+	+	⊗	+	-	⊗
	Cheng Mao, 2022	+	+	+	-	+	+	+	+	+	-	⊗
	Chhaya, 2019	+	+	+	-	+	+	+	+	+	-	⊗
	Detloff, 2016	+	+	+	-	-	-	+	+	+	-	⊗
	Dugan, 2015	+	+	+	-	+	+	+	+	+	-	⊗
	Dugan, 2020	+	+	+	-	+	+	+	+	+	-	⊗
	Dugan, 2021	+	+	+	-	+	+	+	+	+	-	⊗
	Hutchinson, 2004	+	+	+	-	+	-	-	⊗	+	-	⊗
	Li, 2020	+	+	+	-	+	-	+	⊗	+	-	⊗
	Nees, 2016	+	+	+	-	-	-	-	⊗	+	-	⊗
	Sliwinsky, 2018	+	+	+	-	+	+	+	+	+	-	⊗
	Tashiro, 2018	+	+	+	-	-	-	+	+	+	-	⊗
	Ward, 2014	+	+	+	-	+	+	+	+	+	-	⊗
Ward, 2016	+	+	+	-	+	+	+	+	+	-	⊗	

D1: Item 1: Was the allocation sequence adequately generated and applied?
 D2: Item 2: Were the groups similar at baseline or were they adjusted for confounders in the analysis?
 D3: Item 3: Was the allocation adequately concealed?
 D4: Item 4: Were the animals randomly housed during the experiment?
 D5: Item 5: Were the caregivers and/or investigators blinded from knowledge which intervention each animal received during the experiment?
 D6: Item 6: Were animals selected at random for outcome assessment?
 D7: Item 7: Was the outcome assessor blinded?
 D8: Item 8: Were incomplete outcome data adequately addressed?
 D9: Item 9: Are reports of the study free of selective outcome reporting?
 D10: Item 10: Was the study apparently free of other problems that could result in high risk of bias?

Judgement
 ⊗ High
 ⊕ Unclear
 + Low



Supplementary Material 2. Risk of bias assessment of the included preclinical studies according to the SYRCLE’s risk of bias assessment tool.