

Supplementary Table S1. Results of leave-one-out sensitivity analysis for VAS score. A change in the *p*-value indicates that the excluded study significantly influences the observed effect size.

Study removed	Statistics with study removed		
	Mean difference [95% CI]	<i>p</i> -value	<i>I</i> ²
Benito-Ruiz <i>et al.</i> 2009	-16.13 [-25.49, -6.77]	0.0007	89%
Costa <i>et al.</i> 2020	-15.38 [-23.71, -7.05]	0.0003	90%
Kumar <i>et al.</i> 2015 A	-10.07 [-15.76, -4.37]	0.0005	79%
Kumar <i>et al.</i> 2015 B	-10.90 [-17.35, -4.44]	0.0009	85%
Lugo <i>et al.</i> 2016	-15.97 [-25.86, -6.08]	0.002	89%
Luo <i>et al.</i> 2022	-13.90 [-22.18, -5.62]	0.001	90%

CI: confidence interval.

Supplementary Table S2. Results of leave-one-out sensitivity analysis for the total WOMAC score. A change in the *p*-value indicates that the excluded study significantly influences the observed effect size.

Study removed	Statistics with study removed		
	Mean difference [95% CI]	<i>p</i> -value	<i>I</i> ²
Benito-Ruiz <i>et al.</i> 2009	-6.99 [-10.41, -3.57]	< 0.0001	77%
Chen <i>et al.</i> 2023	-7.21 [-10.45, -3.97]	< 0.0001	75%
Costa <i>et al.</i> 2020	-6.51 [-9.80, -3.22]	0.0001	77%
Costa <i>et al.</i> 2021	-6.08 [-9.18, -2.97]	0.0001	76%
Hewling <i>et al.</i> 2019	-7.09 [-10.34, -3.84]	< 0.0001	76%
Jiang <i>et al.</i> 2014	-7.18 [-10.96, -3.39]	0.0002	75%
Kumar <i>et al.</i> 2015 A	-5.68 [-8.68, -2.68]	0.0002	72%
Kumar <i>et al.</i> 2015 B	-5.67 [-8.57, -2.76]	0.0001	72%
Lugo <i>et al.</i> 2016	-6.66 [-10.21, -3.12]	0.0002	77%
Luo <i>et al.</i> 2022	-5.56 [-8.45, -2.66]	0.0002	71%
McAlindon <i>et al.</i> 2011	-7.15 [-10.29, -4.00]	< 0.0001	76%
Schauss <i>et al.</i> 2012	-6.12 [-9.33, -2.90]	0.0002	75%

CI: confidence interval.

Supplementary Table S3. Report of adverse events.

Author	n	Intervention arms	Events, n n (% of AE)	Headache, n (% of AE)	Gastrointestinal, n (% of AE)	Respiratory infections, n (% of AE)	Chest pain, n (% of AE)	Dermatologic, n (% of AE)	Dehydration, n (% of AE)	Sleep pattern disturbance, n (% of AE)	Neurological, n (% of AE)	Allergic reaction, n (% of AE)	Genitourinary, n (% of AE)
Benito-Ruiz <i>et al.</i> 2009	111 96	HC Placebo	50 31	9 (18.0) 5 (16.1)	17 (34.0) 12 (38.7)	5 (10.0) 5 (16.2)	ND ND	ND ND	ND ND	ND ND	ND ND	ND ND	ND ND
Chen <i>et al.</i> 2023	38 37 38 38	HC-II EC + HC-II GS HCl Placebo	ND	ND	ND	ND	ND	ND	ND	ND	ND	ND	ND
Costa <i>et al.</i> 2020	20 20 20	UC-II + physiotherapy Placebo + physiotherapy Placebo + physiotherapy	ND	ND	ND	ND	ND	ND	ND	ND	ND	ND	ND
Costa <i>et al.</i> 2021	20 20 20 20	UC-II Placebo UC-II + physiotherapy Placebo + physiotherapy	ND	ND	ND	ND	ND	ND	ND	ND	ND	ND	ND
Hewling <i>et al.</i> 2019	44 44	Native collagen Placebo	17 6	(4.6) ND	(9.2) (6.9)	(13.7) ND	(2.3) ND	(2.3) ND	(2.3) ND	(2.3) ND	(2.3) ND	(2.3) ND	
Jiang <i>et al.</i> 2014	46 48	BCP Placebo	ND	ND	ND	ND	ND	ND	ND	ND	ND	ND	ND
Kumar <i>et al.</i> 2015	19 11 19 11	PCP Placebo BCP Placebo	1* ND 3* ND	ND ND + ND	ND ND ND ND	ND ND ND ND	ND ND ND ND	ND ND ND ND	ND ND ND ND	ND ND ND ND	ND ND ND ND		
Lugo <i>et al.</i> 2016	63 65 58	UC-II CS Placebo	8 28 9	ND 4 (14.3) 2 (22.2)	3 (37.5) 9 (32.1) 3 (33.3)	2 (25) 4 (14.3) 2 (22.2)	ND 2 (7.1) ND	1 (12.5) 2 (7.1) ND	ND ND ND	ND 1 (3.6) ND	ND ND ND	ND ND ND	
Luo <i>et al.</i> 2022	34 33 34	UC-II GS + CS Placebo	1 ND 2	ND ND ND	ND ND 1 (50)	ND ND ND	ND ND 1 (50)	1 (100) 2 (7.1) ND	ND ND ND	ND ND ND	ND ND ND	ND ND ND	
McAlindon <i>et al.</i> 2011	15 15	HC Placebo	43 45	ND ND	ND ND	ND ND	ND ND	ND ND	ND ND	ND ND	ND ND	ND ND	
Schauss <i>et al.</i> 2012	35 33	HC-II Placebo	3 6	ND ND	ND 3 (50)	ND ND	ND ND	1 (33.3) ND	ND ND	ND ND	ND ND	ND ND	

*Subjects. Adverse events such as pain were not included.
 HC-II: hydrolysed type II collagen; EC: BAND'S essence of chicken; PCP: porcine collagen peptides; BCP: bovine collagen peptides; UC-II: undematured type II collagen; CS: chondroitin sulfate; GS: glucosamine; ND: no data.