

Tables

Table 1. Characteristics of trials included in OA Trial Bank for IA glucocorticosteroid injections

	N at baseline	Type of OA	Glucocorticosteroid intervention	Control interventions	Outcomes	Inflammation	Follow-up
Arden et al. 2008	150	Knee	40mg triamcinolone acetonide and 2ml lignocaine (n=79)	1. Tidal irrigation (n=71)	- Pain (VAS) - WOMAC pain - WOMAC physical functioning - WOMAC stiffness - WOMAC total - Global assessment (5 pt Likert)	Presence of effusion by physical examination (small/moderate/large)	2, 4, 12 and 26 weeks
Atchia et al. 2011	77	Hip	Methylprednisolone acetate (depomedrone) 3ml/120mg (n=19)	1. Placebo (3mg saline) (n=19) 2. Standard care (n=20) 3. Hyaluronic acid (durolane) 3ml/60mg (n=19)	- Pain, worst (NRS) - WOMAC pain - WOMAC physical functioning - WOMAC stiffness	Presence of synovitis >7 mm on ultrasound	1, 4, 8 and 16 weeks
Boon et al. 2010	60	Knee	Methylprednisolone acetate 40mg (n=20)	1. Low dose Botulinum toxin type A (100 units) (n=20) 2. High dose Botulinum toxin type A (200 units) (n=20)	- Pain (VAS) - WOMAC pain - WOMAC physical functioning - WOMAC stiffness - WOMAC total	Presence of effusion by physical examination (mild/moderate/large)	8, 12 and 26 weeks
Chao et al. 2010	79	Knee	40 mg triamcinolone acetonide (n=40)	Placebo (1 cc 0.9% saline) (n=39)	- Pain (VAS) - WOMAC pain - WOMAC total	Pathologic effusion of ≥ 5 mm present on ultrasound	4 and 12 weeks
De Campos et al. 2012	104	Knee	20 mg triamcinolone hexacetonide + 6 mL hylan GF20 (n=52)	Hylan GF20 (6 mL) (n=52)	- Pain (VAS) - WOMAC pain - WOMAC total	-	1, 4, 12 and 24 weeks
Lambert et al. 2007	52	Hip	10 mg bipuvicaine, 40 mg triamcinolone (n=31)	Placebo (10 mg bipuvicaine, 2 ml saline) (n=21)	- WOMAC pain - WOMAC physical functioning - WOMAC stiffness	-	1, 2, 3 and 6 months

					- Global assessment		
Ravaud et al. 1999	98	Knee	3.75 mg cortivazol in 1.5 ml (n=25)	1. Placebo (1.5 ml 0.9% saline) (n=28) 2. joint lavage and IA placebo (n=21) 3. Joint lavage and IA corticosteroid (n=24)	- Pain (VAS) - Global status (VAS)	Evidence of effusion by clinical assessment (present or not)	1, 4, 12 and 24 weeks

Table 2. Baseline characteristics of patients in the study, means (SD) (unless otherwise stated)

	Total population N=620	Comparison 1: Glucocorticosteriod versus placebo N=222	Comparison 2: Glucocorticosteriod versus Hyaluronic acids N=142	Comparison 3: Glucocorticosteriod versus Joint lavage N=196
Age (years)	64.74 (10.47)	64.47 (11.23)	64.15 (10.24)	66.28 (9.56)
Gender, % female	308 (49.7%)	91 (41%)	99 (69.7%)	82 (41.8%)
BMI (kg/m2)	29.82 (5.06)^	28.44 (4.89)*	29.16 (4.73)	30.80 (5.09)
Hip OA, %	129 (20.8%)	90 (40.5%)	38 (26.8%)	-
Knee OA, %	491 (79.2%)	132 (59.5%)	104 (73.2%)	196 (100%)
KL grade, %				
1	35 (5.6%)	10 (4.5%)	20 (14.1%)	4 (2.0%)
2	205 (33.1%)	41 (18.5%)	41 (28.9%)	107 (54.6%)
3	202 (32.6%)	66 (29.7%)	55 (38.7%)	39 (19.9%)
4	71 (11.5%)	25 (11.3%)	25 (17.6%)	20 (10.2%)
missing	107 (17.3%)	80 (64.0%)**	1 (0.7%)	26 (13.3%)
Duration of complaints (months)	76.98 (102.57^^	86.19 (122.62)	40.64 (36.02)##	69.36 (87.22)
Inflammation, %	259 (41.8%)^^^	88 (39.6%)**	28 (19.7%)#	110 (56.1%)
Severe pain (≥70 points), %	213 (34.4%)	61 (27.5%)	60 (42.3%)	65 (33.2%)\$\$
Pain (0-100)	59.92 (20.34)	58.63 (17.58)	64.50 (21.21)	57.71 (23.02)\$

^N=485; ^^ N=495; ^^N=458; * N=91 (Not available for Lambert and Chao); ** Not available for Chao; ***N=165 (Not available for Lambert)

N=38 (Not available for de Campos); ## N=36 (not available for de Campos)

\$ N=187; \$\$ N=187

Table 3. Risk of Bias assessment

	A1	B2	C3	C4	C5	D6	D7	E8	E9	E10	E11
Arden et al. 2008	+	?	-	-	-	+	?	-	+	+	+
Atchia et al. 2011	+	?	-	-	-	+	+	-	?	+	+
Boon et al. 2010	+	?	+	?	+	-	?	-	+	+	+
Chao et al. 2010	+	?	+	?	+	-	?	+	?	+	+
de Campos et al. 2012	+	?	+	?	+	+	?	+	?	+	+
Lambert et al. 2007	+	+	+	+	+	+	+	-	+	+	+
Ravaud et al. 1999	+	?	+	+	+	+	+	+	+	+	+

A1. method of randomization adequate; B2. Treatment allocation concealed; C3. patient blinded to the intervention; C4. care provider blinded to the intervention; C5. outcome assessor blinded to the intervention; D6. drop-out rate described and acceptable; D7; randomized participants analysed in the group to which they were allocated; E8. groups similar at baseline regarding the most important prognostic indicators; E9. co-interventions avoided or similar; E10. compliance acceptable; E11. timing of the outcome assessment similar in all groups.

Table 4. Overall effectiveness on primary outcome pain severity (0-100 scale)

	N total	N intervention group	N control group	Effect estimate (95% CI)	Adjusted effect estimate* (95% CI)	I ²	p-value
Short term (4 trials)¹	207	107	100	17.74 (11.65;23.82)	18.72 (13.04;24.41)	67%	<0.001
Mid-term (4 trials)¹	181	98	83	7.99 (1.44;14.53)	10.00 (3.88;16.13)	67%	0.002
Long term (2 trials)²	71	42	29	6.54 (-6.94;20.02)	6.25 (-8.59;21.10)	0%	0.403
Short term (2 trials)³	142	71	71	9.06 (5.05;13.08)	9.38 (5.69;13.09)	0%	<0.001
Mid-term (2 trials)³	131	66	65	0.66 (-3.46;4.77)	0.97 (-2.96;4.90)	0%	0.627
Long term (1 trial)⁴	93	47	46	-0.38 (-5.01;4.25)	0.22 (-4.30;4.75)	n.a.	0.921
Short term (2 trials)⁵	191	102	89	0.83 (-1.63;3.30)	1.08 (-1.31; 3.47)	51%	0.374
Mid-term (2 trials)⁵	179	93	86	-1.60 (-4.36;1.16)	-1.21 (-3.83;1.41)	51%	0.363
Long term (2 trials)⁵	172	90	82	-4.85 (-7.68;-2.02)	-4.57 (-7.40;-1.74)	51%	0.02

* adjusted for baseline pain, age and gender; statistical significant differences (p<0.05) in bold; n.a. Not Applicable

¹Atchia et al. 2011, Chao et al. 2010,

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Atchia et al. 2011,

de Campos et al. 2012; ⁴ de Campos et al. 2012; ⁵ Arden et al. 2008,

Table 5. Interaction effects of severe pain (≥ 70 points) with IA glucocorticoid injection for primary outcome pain severity (0-100 scale)

	Severe pain n/ N glucocorticosteroid group	Severe pain n/ N control group	Unadjusted interaction effect estimate (95%CI)	Adjusted interaction effect estimate# (95%CI)	Effect Size	Adjusted p-value for interaction term
IA glucocorticosteroid injection versus placebo (4 trials)						
Short term pain ¹	28/107	30/100	14.93 (2.24;27.63)	13.91 (1.50;26.31)	0.56	0.028
Mid-term pain ¹	26/98	19/83	3.01 (-10.66;16.87)	1.84 (-11.27;14.94)	0.07	0.782
Long term pain ²	15/42	11/29	-4.23 (-32.56;24.09)	-4.31 (-33.73;25.11)	-0.17	0.771
IA glucocorticosteroid injection versus Hyaluronic acid (2 trials)						
Short term pain ³	33/71	27/71	0.85 (-7.05;8.76)	3.11 (-4.36;10.59)	0.10	0.412
Mid-term pain ³	31/47	25/65	-4.68 (-12.82;3.45)	-3.15 (-11.14;4.85)	-0.10	0.438
IA glucocorticosteroid injection versus tidal irrigation (2 trials)						
Short term pain ⁴	32/96	33/87	1.85 (-3.19;6.90)	1.28 (-3.68;6.24)	0.04	0.610
Mid-term pain ⁴	29/88	28/83	0.30 (-5.44;6.05)	-0.28 (-5.77;5.21)	-0.01	0.919
Long term pain ⁴	28/85	27/79	4.12 (-1.82;10.06)	3.42 (-2.44;9.29)	0.10	0.251

adjusted for age and gender and baseline pain

¹Atchia et al. 2011, Chao et al. 2010,
de Campos et al. 2012; ⁴ Arden et al. 2008,

Table 6. Interaction effects of inflammation with IA glucocorticoid injection for primary outcome pain severity (0-100 scale)

	Inflammation n/ N intervention group	Inflammation n/N control group	Unadjusted interaction effect estimate (95%CI)	Adjusted interaction effect estimate# (95%CI)	Effect Size	Adjusted p- value for interaction term
<i>IA glucocorticosteriod versus placebo (3 trials)</i>						
Short term pain ¹	39/77	43/79	2.14 (-11.36;15.65)	7.34 (-4.97;19.65)	0.29	0.24
Mid-term pain ¹	39/71	34/66	3.76 (-10.52;18.04)	11.08 (-1.34;23.50)	0.44	0.08
<i>IA glucocorticosteriod versus tidal irrigation (2 trials)</i>						
Short term pain ²	60/102	47/89	1.29 (-3.68;6.25)	0.73 (-4.07;5.53)	0.02	0.765
Mid-term pain ²	58/93	46/86	1.68 (-3.94;7.30)	1.77 (-3.47;7.01)	0.05	0.505
Long term pain ²	57/90	43/82	4.25 (-1.50;10.00)	5.24 (-0.35;10.84)	0.16	0.066

adjusted for age and gender and baseline pain

¹Atchia et al. 2011, Chao et al. 2010,

Arden et al. 2008,

Table 7. Interaction effects of inflammation measured by ultrasound with IA glucocorticoid injection for primary outcome pain severity (0-100 scale)

	Total N	Inflammation n/ N intervention group	Inflammation n/ N control group	Unadjusted interaction effect estimate (95%CI)	Adjusted interaction effect estimate# (95%CI)	Effect size	Adjusted p- value for interaction term
<i>IA glucocorticosteroid versus placebo (2 trials)</i>							
Short term pain	103	29/52	27/51	9.22 (-5.21;23.64)	9.04 (-0.71;18.80) ¹	0.39	0.069
Mid-term pain	96	29/49	25/47	6.53 (-9.18;22.24)	7.14 (-4.27;18.55) ²	0.31	0.217

adjusted for age and gender and baseline pain

¹ Baseline pain: 0.84 (0.69;0.99), p<0.001

² Baseline pain: 0.82 (0.65;0.99), p<0.001

¹Atchia et al. 2011, Chao et al. 2010

