

## **The health effects of a low-inflammatory diet in adults with arthritis: systematic review and meta-analysis**

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### **Supplementary Tables, Figures and Search Syntax:**

*Supplementary Table 1: Table illustrating which measures were multiplied by -1*

PROM	Low Score	High score	Multiply by -1?
<b>Physical Health:</b>			
- HAQ		Poor Health	No
- AIMS2 Physical		Poor Health	No
- SF-36 Physical	Poor Health		Yes
<b>General Health:</b>			
- VAS Global	Poor Health		No
- VAS Health	Poor Health		No
- SF-36 Global Health	Poor Health		No
<b>Pain Scores:</b>			
- SF-36 bodily pain	Poor Health		Yes
- Pain VAS		Poor Health	No
- AIMS2 Symptom		Poor Health	No
- ICOAP Total		Poor Health	No

*Supplementary Table 2: Risk of Bias assessment for non-randomised trials utilising Cochrane's ROBINS-I tool*

Domain	Adam et. al. 2003	McKellar et. al. 2007
Bias due to confounding	Low risk	Serious risk
Bias in selection of participants into the study	Low risk	Low risk
Bias in classification of interventions	Low risk	Moderate risk
Bias due to deviations from intended interventions	Moderate risk	Low risk
Bias due to missing data	Moderate risk	Low risk
Bias in measurement of outcomes	Moderate risk	Moderate risk
Bias in selection of the reported result	Moderate risk	Low risk
Overall	<i>Moderate risk</i>	<i>Serious risk</i>

*Supplementary Table 3: Assessing risk of bias in the dietary interventions implemented in the studies.*

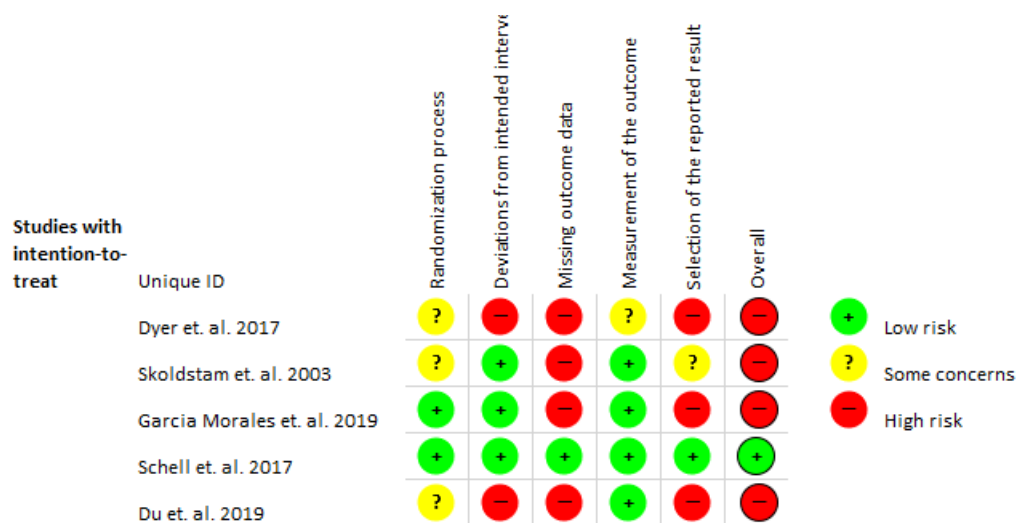
<u>Author and Year</u>	<u>Was a dietitian involved in the designing of the dietary intervention ?</u>	<u>Was the dietary intervention a partial or full simulation of the low inflammatory/Mediterranean diet?</u>	<u>Was there monitoring of adherence to the dietary intervention?</u>	<u>Was a validated tool utilised to measure dietary adherence?</u>
Dyer et. al., 2017	Not Specified	Full simulation of Mediterranean Diet	Yes, a 7 day food diary (start, middle and end) + compliance score	No - self developed compliance score 0-100
Skoldstam et. al., 2003	Not Specified	Modified Mediterranean for Swedish people.	Yes, 3 week BD meal program, weekly dietician phone and 3rd weekly visits	No, had questionnaires and dietary interviews to assess adherence, but no specific mention of any validated tool utilised.
Garcia Morales et. al., 2019	Yes	Full simulation of Mediterranean Diet	Yes, however used a 24hr food recall questionnaire	No
Schell et. al., 2017	Not Specified	Partial simulation of Low inflammatory diet	Yes - 3 day food diary (week 6, 12, 14, 20, 26)	No
Du et. al., 2019	Not Specified	Partial simulation of Low inflammatory diet	Yes, calendars were given to patients to remind them of consumption, but compliance was based on self reporting.	No
Mckellar et. al., 2007	Not Specified	Partial simulation of Mediterranean Diet (refer to it as "Mediterranean-type diet")	Yes, using FFQ questionnaires	Yes
Adam et. al., 2003	Not Specified	Modified AID (lactoveg)	Yes - monthly counseling with 3 days recorded prior to visit	No

**Supplementary Table 4: Individual study intervention results**

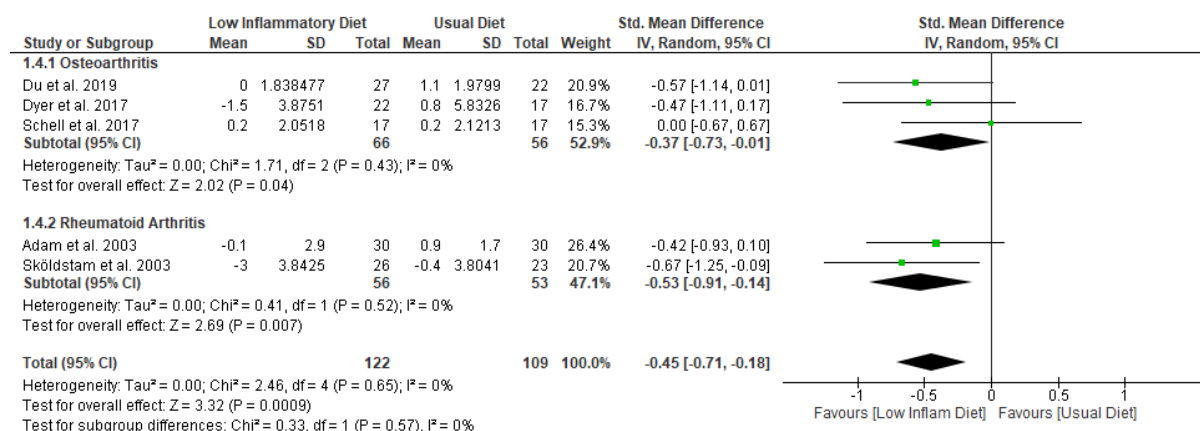
Outcome	Specific Measure	Study	N=	0-2 Months MEAI	0-2 Months SCI	2-4 Months MEAI	2-4 Months SCI	4-6 months MEAI	4-6 months SCI
Weight Change	Kg	Adam 2003	30			-0.1		2.9	
		Garcia Morales 2019	35						
		Skoldstam 2003	26			-3		3.8425	
		Dyer 2017	22			-1.5		3.8751	
Weight Change	BMI	Du 2019	27			0		1.838477	
		Schell 2017	17			0.2		2.0518	
Change in Inflammatory Markers	ESR	Skoldstam 2003	26	7	5.3852	1		4.1602	
		(hs)/CRP	Skoldstam 2003	24 (0-2m) / 25 (2-4m)	10	11.9181	-5		5
	IL-6	Schell 2017	17			-1.1		1.5	
		Schell 2017	17			-5.4		0.6403	
	IL-18	Dyer 2017	29			-1.76		1.1463	
		Schell 2017	17			-11.1		4.0608	
	Dyer 2017	29			-0.47		0.3666		
PROM	HAQ(-DI) - (measure functional status in RA, OA, etc.)	Skoldstam 2003	26 (n=25 for 0-2m)	-0.1	0.1401	-0.1		0.1266	
		Garcia Morales 2019	35					0.55	0.6
		Schell 2017	17			-0.2		0.1414	
	AIMS2 Physical (subscore assessing physical function - high score equals poor health - can correlate with HAQ scores)	Dyer 2017	49	0	0.3239	-0.1		0.2931	
		SF-36 physical function	Skoldstam 2003	26			-2.5		15.2
	DAS28 (Measure of RA disease activity)	Garcia Morales 2019	35						
		Skoldstam 2003	26	-0.2	0.3616	-0.5		0.3328	
	VAS Global VAS Health	Skoldstam 2003	26	-4	5.9646	-12		5.0115	
		Schell 2017	17			0.3		0.2236	
	SF-36 Global Health	Skoldstam 2003	26			5.7		14.6	
		Garcia Morales 2019	35					-0.5	25.7
	SF-36 Bodily Pain	Skoldstam 2003	26			-4.5		24.3	
		Garcia Morales 2019	35					-4.2	24.3
	Pain VAS	Skoldstam 2003	26	-2	5.831	-12		4.6781	
		Schell 2017	17			-0.6		0.2236	
AIMS2 Symptom (Pain) ICOAP (Total Pain)	Dyer 2017	49	-0.1	0.5459	-0.3		0.5153		
	Schell 2017	17			-16		4.827		
ROM	Knee Flexion	Dyer 2017	33			10		5.2223	
		Dyer 2017	33			14		4.324	
		Dyer 2017	33			5		3.8925	

Supplementary table 5: Individual study control/usual diet results

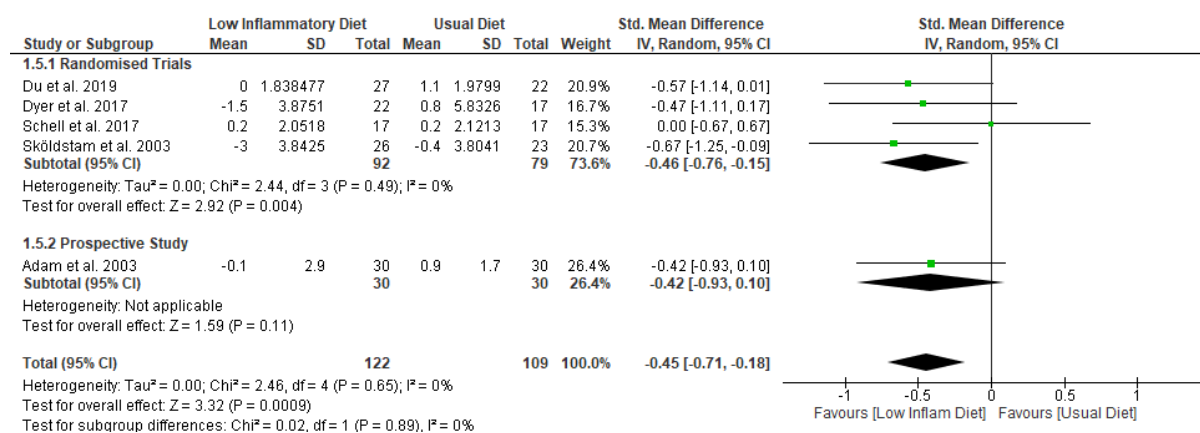
Outcome	Specific Measures	Study	N=	0-2 Months MEAN	0-2 Months SD	2-4 Months MEAN	2-4 Months SD	4-6 months MEAN	4-6 months SD		
Weight Change	Kg	Adam 2003	30			0.9		1.7			
		Garcia Morales 2019	27								
		Skoldstam 2003	23			-0.4		3.8041		-0.8	2.4591
		Dyer 2017	17			0.8		5.8326			
		Du 2019	22			1.1		1.9799			
Weight Change	BMI	Schell 2017	17			0.2		2.1213			
		Skoldstam 2003	25		-1	4.2426		2		4.8415	
Change in Inflammatory Markers	ESR	Skoldstam 2003	25								
		(hs)/CRP	Skoldstam 2003	23		-3	3.4704		0		3.8448
		Schell 2017	17			-0.9		1.4422			
	IL-6	Schell 2017	17			-0.1		1.456			
	IL-18	Schell 2017	17			-0.22		0.4115			
		Schell 2017	17			-2.4		4.1761			
		Dyer 2017	25			0.19		0.4136			
PROM	HAQ(-DI) - (measure functional status in RA, OA, etc.)	Skoldstam 2003	23		-0.1	0.1769		0		0.1769	
		Garcia Morales 2019	27						0.82	0.68	
		Schell 2017	17				0	0.1414			
	AIMS2 Physical (subscore assessing physical function - high score equals poor health - can correlate with HAQ scores)	Dyer 2017	49		-0.1	0.3839		-0.1	0.3839		
	SF-36 physical function	Skoldstam 2003	25				-1.4	13.4			
		Garcia Morales 2019	27						0	17.3	
	DAS28 (Measure of RA disease activity)	Skoldstam 2003	23		-0.1	0.4128		0	0.4278		
	VAS Global	Skoldstam 2003	25		-2	5.8		-1	5.8		
	VAS Health	Schell 2017	17				0	0.1414			
	SF-36 Global Health	Skoldstam 2003	25				0.7	21.7			
		Garcia Morales 2019	27						-7.8	18.8	
	SF-36 Bodily Pain	Skoldstam 2003	25				-4	20.1			
		Garcia Morales 2019	27						4.4	30.8	
	Pain VAS	Skoldstam 2003	25		2	6.2482		3	5.8		
		Schell 2017	17				-0.4	0.2828			
AIMS2 Symptom (Pain)	Dyer 2017	49		-0.6	0.5459		-0.6	0.5557			
ICOAP (Total Pain)	Schell 2017	17				-5.5	4.3139				
ROM	Knee Flexion	Dyer 2017	28					-5	6.2593		
	Hip Rotation	Dyer 2017	28					-1	5.3486		
	Hip Flexion	Dyer 2017	28					1	2.8094		



Supplementary Figure 1: Risk of Bias assessment for randomised trials using Cochrane's ROB-V2 methodology

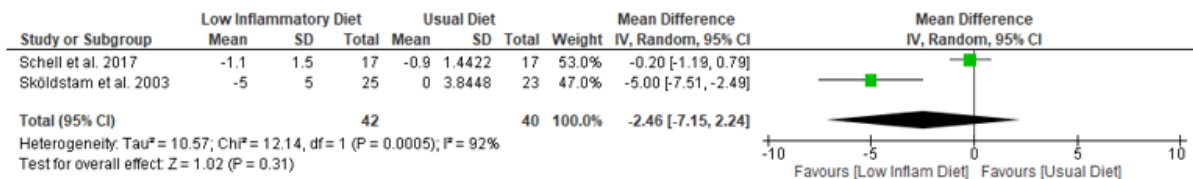


Supplementary Figure 2: Subgroup analysis of weight change by diagnosis following 2-4 months of intervention/control

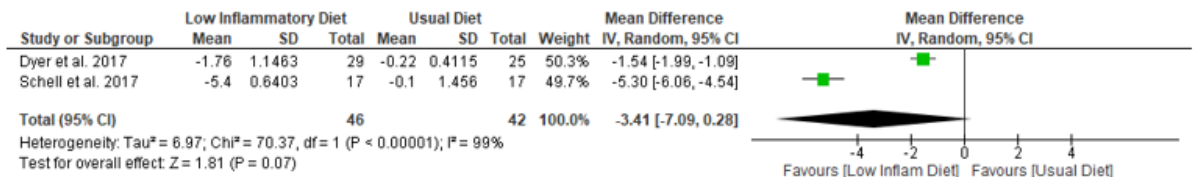


Supplementary Figure 3: Subgroup analysis of weight change by study type following 2-4 months of intervention/control

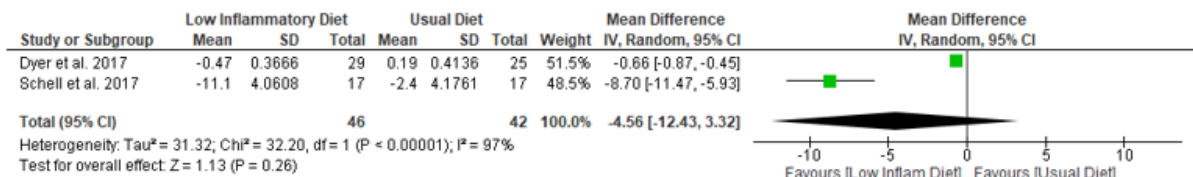
**A) Change in CRP at 2-4 months**



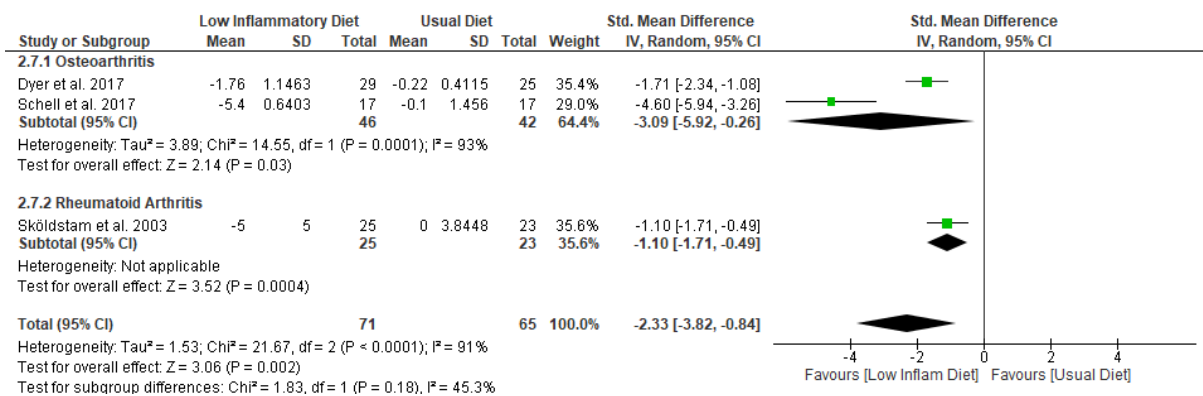
**B) Change in IL-6 at 2-4 months**



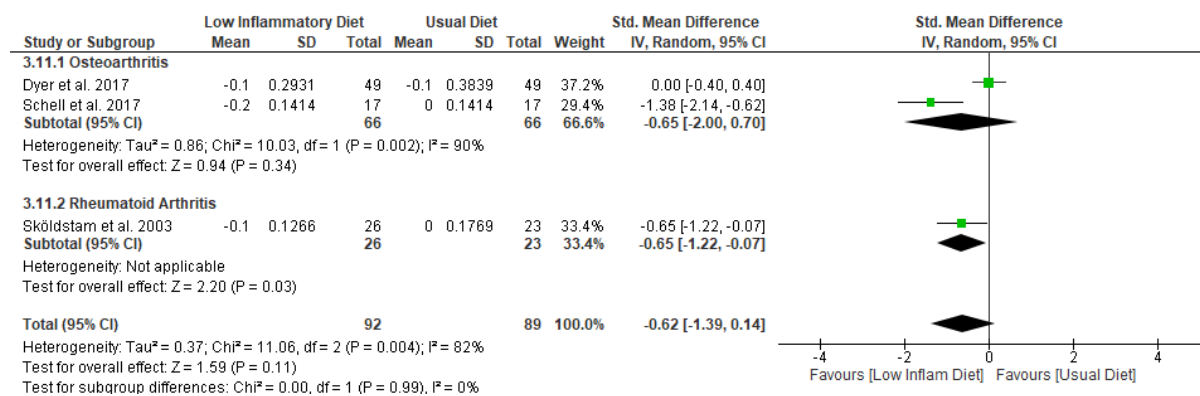
**C) Change in IL-1β at 2-4 months**



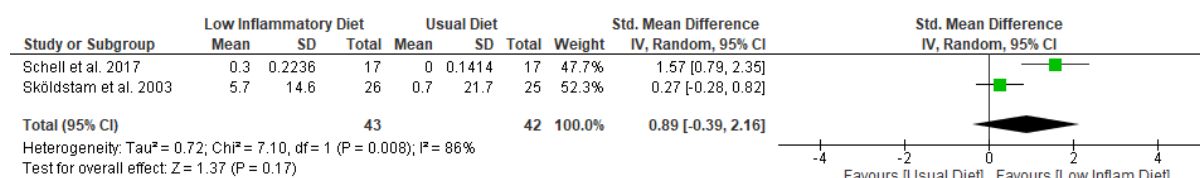
Supplementary Figure 4: Meta-analysis of change in specific inflammatory biomarkers after 2-4 months of intervention/control.



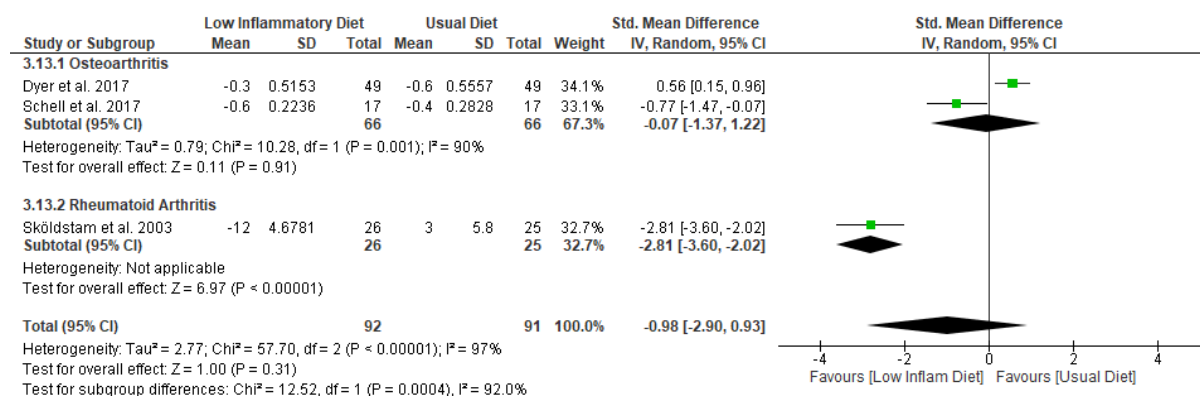
Supplementary Figure 5: Meta-analysis of change in any inflammatory biomarker by diagnosis after 2-4 months of intervention/control.



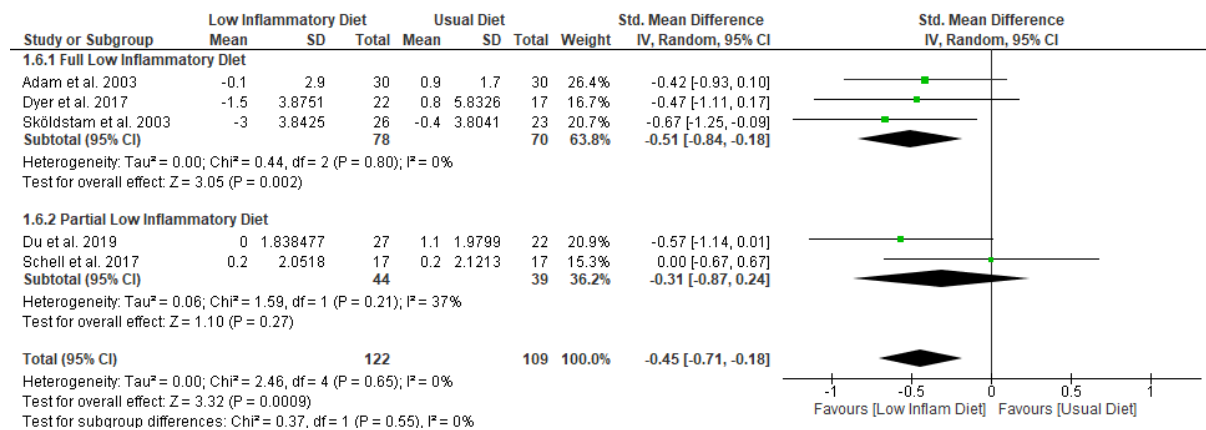
*Supplementary Figure 6: Meta-analysis of change in any physical function outcomes by diagnosis after 2-4 months of intervention/control.*



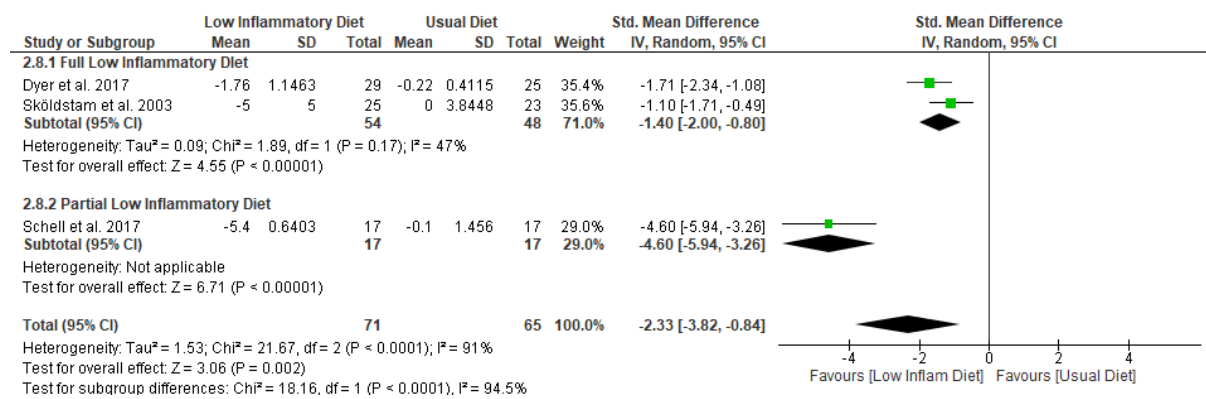
*Supplementary Figure 7: Meta-analysis of change in any general health measure after 2-4 months of intervention/control.*



*Supplementary Figure 8: Meta-analysis of change in any pain scores by diagnosis after 2-4 months of intervention/control.*

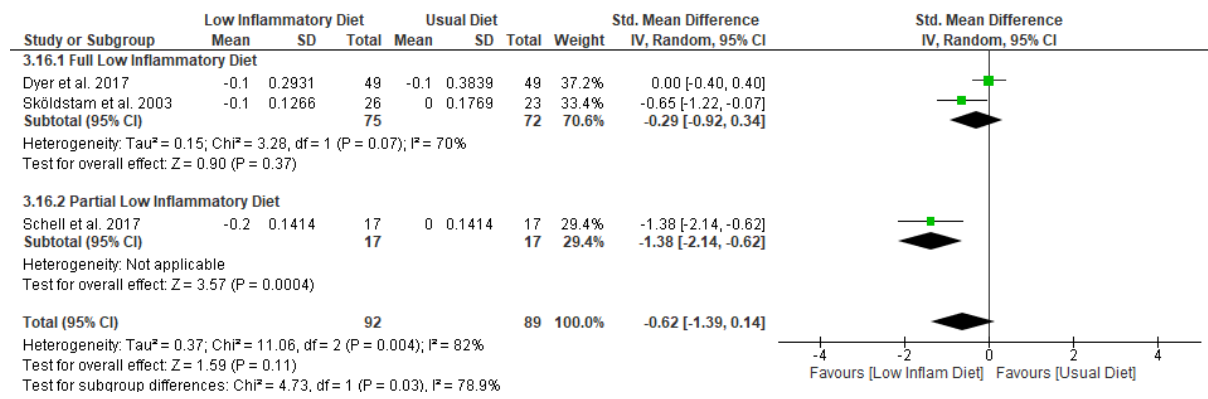


*Supplementary Figure 9: Subgroup analysis of weight change by nature of intervention (full vs partial low-inflammatory diet) following 2-4 months of intervention/control.*

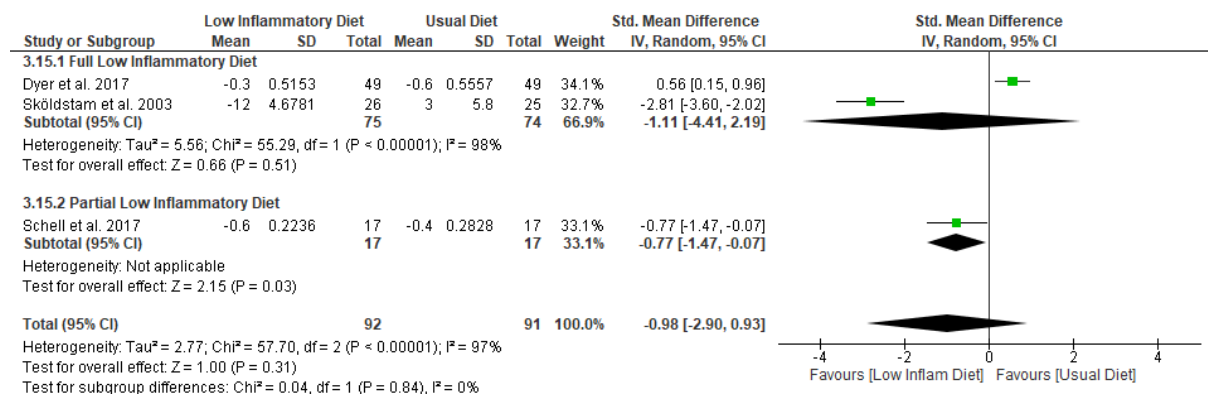


*Supplementary Figure 10: Meta-analysis of change in any inflammatory biomarker nature of intervention (full vs partial low-inflammatory diet) after 2-4 months of intervention/control (Note: A meta-analysis comparing interventions that resulted in weight loss compared to no weight loss produced the same analysis. Weight loss included the same trials as full low-inflammatory diet, no weight loss included the same trials as partial low-inflammatory diet).*

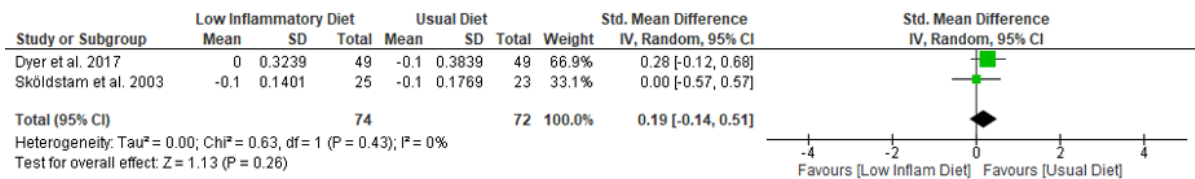




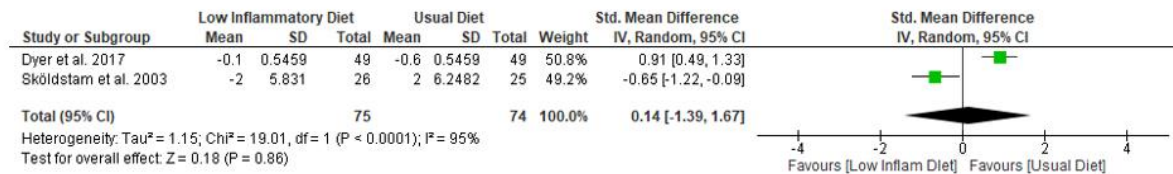
*Supplementary Figure 11: Meta-analysis of change in any physical function outcomes by nature of intervention (full vs partial low-inflammatory diet) after 2-4 months of intervention/control. (Note: A meta-analysis comparing interventions that resulted in weight loss compared to no weight loss produced the same analysis. Weight loss included the same trials as full low-inflammatory diet, no weight loss included the same trials as partial low-inflammatory diet).*



*Supplementary Figure 12: Meta-analysis of change in any pain scores by nature of intervention (full vs partial low-inflammatory diet) after 2-4 months of intervention/control. (Note: A meta-analysis comparing interventions that resulted in weight loss compared to no weight loss produced the same analysis. Weight loss included the same trials as full low-inflammatory diet, no weight loss included the same trials as partial low-inflammatory diet).*



*Supplementary Figure 13: Meta-analysis of physical function change when comparing low inflammatory diet to usual diet following 0-2 months of the intervention/control.*



*Supplementary Figure 14: Meta-analysis of pain score change when comparing low inflammatory diet to usual diet following 0-2 months of the intervention/control.*

**SEARCH SYNTAX:**

A. Syntax for MEDLINE, EMBASE, CENTRAL and Cochrane Databases for Systematic Reviews (note: this syntax would be used using the OvidSP Platform searching through the four databases):

1. arthritis/
2. arthritis, psoriatic/
3. exp arthritis, rheumatoid/
4. exp osteoarthritis/
5. (arthritis adj5 (psoriatic or psoriasis)).tw.
6. (arthritis adj5 rheumatoid).tw.
7. (arthritis adj5 (seronegative or seropositive)).tw.
8. osteoarthritis.tw.
9. arthritis, reactive/
10. spondylitis, ankylosing/
11. (arthritis adj5 reactive).tw.
12. (spondylitis adj5 ankylosing).tw.
13. (arthritis adj5 IBD-related).tw.
14. (arthritis adj5 IBD-associated).tw.
15. or/1-14
16. limit 15 to "all adult (19 plus years)"
17. (low Inflammatory adj3 diet\*).tw.
18. (low Inflammatory adj3 dietary pattern\*).tw.
19. (low Inflammatory adj3 type diet\*).tw.
20. (anti-Inflammatory adj3 diet\*).tw.
21. (anti-Inflammatory adj3 dietary pattern\*).tw.
22. (anti-Inflammatory adj3 type diet\*).tw.
23. Diet, Mediterranean/
24. (mediterranean adj3 diet\*).tw.
25. (mediterranean adj3 dietary pattern\*).tw.
26. (mediterranean adj3 type diet\*).tw.
27. MedDiet.tw.
28. MeDi.tw.
29. MeDiet.tw.
30. ((crete or cretan) adj4 diet\*).tw.
31. animals/ not (animals/ and humans/)
32. or/17-30
33. 16 and 32
34. 33 not 31
35. remove duplicates from 34

## B. Syntax for CINAHL

S1	arthritis
S2	psoriatic arthritis
S3	rheumatoid arthritis
S4	osteoarthritis
S5	arthritis N5 (psoriasis OR psoriatic)
S6	arthritis N5 (rheumatoid)
S7	arthritis N5 (seronegative OR seropositive)
S8	TX osteoarthritis
S9	reactive arthritis
S10	ankylosing spondylitis
S11	arthritis N5 (reactive)
S12	spondylitis N5 (ankylosing)
S13	arthritis N5 (IBD-related)
S14	arthritis N5 (IBD-associated)
S15	S1 OR S2 OR S3 OR S4 OR S5 OR S6 OR S7 OR S8 OR S9 OR S10 OR S11 OR S12 OR S13 OR S14
S16	S1 OR S2 OR S3 OR S4 OR S5 OR S6 OR S7 OR S8 OR S9 OR S10 OR S11 OR S12 OR S13 OR S14 Limiters - Human; Age Groups: Adult: 19-44 years, Middle Aged: 45-64 years, Aged: 65+ years, Aged, 80 and over
S17	low inflammatory N3 (diet)
S18	low inflammatory N3 (dietary pattern)
S19	low inflammatory N3 (type diet)
S20	mediterranean diet
S21	Mediterranean N3 (Diet)
S22	Mediterranean N3 (Dietary pattern)
S23	Mediterranean N3 (type diet)
S24	TX MedDiet
S25	TX MeDi
S26	TX MeDiet
S27	(crete or cretan) N4 diet
S28	Anti-inflammatory N3 (diet)
S29	Anti-inflammatory N3 (dietary pattern)
S30	anti-inflammatory N3 (type diet)
S31	S17 OR S18 OR S19 OR S20 OR S21 OR S22 OR S23 OR S24 OR S25 OR S26 OR S27 OR S28 OR S29 OR S30
S32	S17 OR S18 OR S19 OR S20 OR S21 OR S22 OR S23 OR S24 OR S25 OR S26 OR S27 OR S28 OR S29 OR S30 Limiters - Human
S33	S16 AND S32