

**Table 1. Summary of overall evidence for the association between systemic inflammation and individual symptoms of depression**

DEPRESSIVE SYMPTOM	PREVALENCE (%)	DOMAIN	STRENGTH OF OVERALL EVIDENCE	EFFECT SIZE*	ROBUST TO MULTI-VARIABLE ADJUSTMENT	SUPPORT FOR TEMPORALITY	CONSISTENT ACROSS INFLAMMATORY MARKERS	HETEROGENEITY OF META-ANALYSIS ( $I^2$ , $\tau^2$ )†	GENERALISABLE ACROSS SUBGROUPS
<b>SUPPORT FOR AN ASSOCIATION</b>									
1. Changes in appetite	7.4	Physical symptom	STRONG	LARGE	YES	YES	YES	LOW (0%, 0.0003)	YES
2. Felt everything was an effort	16	Physical symptom	STRONG	LARGE	YES	YES	YES	HIGH (79%, 0.0142)	YES
3. Could not get going/energy	19.6	Physical symptom	STRONG	LARGE	YES	YES	YES	LOW (21%, 0.0023)	YES
4. Little interest in doing things/unmotivated	11.2	Cognitive symptom	MODERATE	LARGE	YES	YES‡	MISSING DATA	LOW (22%, 0.0010)	YES
5. Sleep was restless	21.5	Physical symptom	MODERATE	MODERATE	YES	YES	YES	HIGH (67%, 0.0031)	YES
<b>EVIDENCE AGAINST AN ASSOCIATION</b>									
6. Bothered by things	6.7	Emotional symptom	STRONG	NO ASSOCIATION	NO	NO	YES	LOW (14%, 0.0027)	N/A
7. Felt hopeless about the future	18.2	Emotional symptom	STRONG	NO ASSOCIATION	NO	NO	YES	LOW (0%, 0.0016)	N/A
8. Felt fearful	3.8	Emotional symptom	MODERATE	NO ASSOCIATION	NO	NO	YES	HIGH (61%, 0.0131)	N/A
9. Life had been a failure	3.5	Emotional symptom	MODERATE	NO ASSOCIATION	NO	NO	YES	HIGH (62%, 0.0160)	N/A

Table 1. (continued)

DEPRESSIVE SYMPTOM	PREVALENCE (%)	DOMAIN	STRENGTH OF OVERALL EVIDENCE	EFFECT SIZE*	ROBUST TO MULTI-VARIABLE ADJUSTMENT	SUPPORT FOR TEMPORALITY	CONSISTENT ACROSS INFLAMMATORY MARKERS	HETEROGENEITY OF META-ANALYSIS ( $I^2$ , $\tau^2$ ) <sup>†</sup>	GENERALISABLE ACROSS SUBGROUPS
<b>UNCERTAIN EVIDENCE</b>									
10. Difficulties concentrating	8.7	Cognitive symptom	MODERATE	MODERATE	YES	YES	YES	MODERATE (47%, 0.0031)	NO
11. Could not shake off the blues	4.3	Perception of self	MIXED	MODERATE	NO	NO	NO	LOW (0%, 0.0016)	N/A
12. Felt worse than others	13.6	Perception of self	MIXED	NO ASSOCIATION	NO	YES	NO	LOW (0%, 0.0016)	N/A
13. Felt depressed	10.2	Emotional symptom	MIXED	MODERATE	YES	YES	NO	MODERATE (43%, 0.0026)	N/A
14. Felt unhappy	12.5	Emotional symptom	MIXED	MODERATE	NO	YES	YES	MODERATE (49%, 0.0068)	N/A
15. Talked less than usual	7	Physical symptom	MIXED	NO ASSOCIATION	NO	YES	NO	LOW (0%, < 0.0001)	N/A
16. Felt lonely	11.5	Emotional symptom	MIXED	MODERATE	NO	YES	NO	LOW (0%, 0.0002)	N/A
17. People were unfriendly	3.1	Cognitive symptom	MIXED	NO ASSOCIATION	NO	NO	YES	MODERATE (36%, 0.0189)	N/A
18. Did not enjoy life	10.6	Emotional symptom	MIXED	MODERATE	NO	NO	NO	MODERATE (29%, 0.0028)	N/A
19. Had crying spells	2.5	Physical symptom	MIXED	LARGE	NO	NO	NO	MODERATE (36%, 0.0180)	N/A
20. Felt sad	14.1	Emotional symptom	MIXED	SMALL	NO	NO	NO	LOW (21%, 0.0017)	N/A
21. People dislike me	2.4	Perception of self	MIXED	NO ASSOCIATION	NO	NO	NO	HIGH (57%, 0.0326)	N/A
22. Feeling bad about yourself	5.3	Perception of self	MIXED & LACKING	MODERATE	NO	MISSING DATA	MISSING DATA	LOW (0%, 0.0002)	N/A
23. Moving or speaking slowly/ too fast	3.6	Physical symptom	MIXED & LACKING	MODERATE	NO	MISSING DATA	MISSING DATA	LOW (0%, 0.0015)	N/A

<b>24. Thought you would be better off dead</b>	1.1	Self-harm symptom	MIXED & LACKING	LARGE	NO	MISSING DATA	MISSING DATA	LOW (19%, 0.0095)	N/A
---	-----	-------------------	-----------------	-------	----	--------------	--------------	-------------------	-----

\*Large, odds ratio  $\geq 1.20$  and  $P < 0.05$ ; Moderate, odds ratio between 1.10 and 1.19 and  $P < 0.05$ ; Small, odds ratio  $< 1.10$ , but  $P < 0.05$ ; No association,  $P > 0.05$ .

†Low,  $I^2 < 25\%$ ; Moderate,  $I^2 25\% - 50\%$ ; High,  $I^2 > 50\%$ ; ‡ Results limited to Health and Retirement Study (HRS)