# SUPPLEMENTAL MATERIAL APPENDIX

**Title:** Association between depressive symptoms or depression and health outcomes for low back pain: a systematic review and meta-analysis

Running title: Depression and outcomes for back pain

**Authors:** Jessica J. Wong, DC, MPH; Andrea C. Tricco, MSc, PhD; Pierre Côté, DC, PhD; Catherine Y. Liang, MPH; Jeremy A. Lewis, BA; Zachary Bouck, MPH; Laura C. Rosella, MHSc, PhD

**Corresponding author**: Jessica J. Wong, Epidemiology Division, Dalla Lana School of Public Health, University of Toronto, 155 College Street, 6th Floor, Toronto, Ontario M5T 3M7, Canada; email: jes.wong@mail.utoronto.ca

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# Appendix I. Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) 2020 Checklist

#### Table 1. PRISMA 2020 item checklist

| Section and Topic                | Item # | Checklist item   |                |  |  |
|----------------------------------|--------|--|----------------|--|--|
| TITLE                            |        |  | Page #         |  |  |
| Title                            | 1      | Identify the report as a systematic review.  | Title page     |  |  |
| ABSTRACT                         |        |  |                |  |  |
| Abstract                         | 2      | See the PRISMA 2020 for Abstracts checklist (Table 2).   | 3              |  |  |
| INTRODUCTION                     |        |  |                |  |  |
| Rationale                        | 3      | Describe the rationale for the review in the context of existing knowledge.  | 5-6            |  |  |
| Objectives                       | 4      | Provide an explicit statement of the objective(s) or question(s) the review addresses.   | 6              |  |  |
| METHODS                          |        |  |                |  |  |
| Eligibility criteria             | 5      | Specify the inclusion and exclusion criteria for the review and how studies were grouped for the syntheses.  | 7-8            |  |  |
| Information sources              | 6      | Specify all databases, registers, websites, organisations, reference lists and other sources searched or consulted to identify studies. Specify the date when each source was last searched or consulted.  | 8              |  |  |
| Search strategy                  | 7      | Present the full search strategies for all databases, registers and websites, including any filters and limits used.   | 8, Appendix    |  |  |
| Selection process                | 8      | Specify the methods used to decide whether a study met the inclusion criteria of the review, including how many reviewers screened each record and each report retrieved, whether they worked independently, and if applicable, details of automation tools used in the process.                     |                |  |  |
| Data collection process          | 9      | Specify the methods used to collect data from reports, including how many reviewers collected data from each report, whether they worked independently, any processes for obtaining or confirming data from study investigators, and if applicable, details of automation tools used in the process. | 9              |  |  |
| Data items                       | 10a    | List and define all outcomes for which data were sought. Specify whether all results that were compatible with each outcome domain in each study were sought (e.g. for all measures, time points, analyses), and if not, the methods used to decide which results to collect.                        | 9              |  |  |
|                                  | 10b    | List and define all other variables for which data were sought (e.g. participant and intervention characteristics, funding sources). Describe any<br>assumptions made about any missing or unclear information.  | 9              |  |  |
| Study risk of bias<br>assessment | 11     | Specify the methods used to assess risk of bias in the included studies, including details of the tool(s) used, how many reviewers assessed each<br>study and whether they worked independently, and if applicable, details of automation tools used in the process.                                 | 9-10           |  |  |
| Effect measures                  | 12     | Specify for each outcome the effect measure(s) (e.g. risk ratio, mean difference) used in the synthesis or presentation of results.  | 10-11          |  |  |
| Synthesis methods                | 13a    | Describe the processes used to decide which studies were eligible for each synthesis.  | 10-11          |  |  |
|                                  | 13b    | Describe any methods required to prepare the data for presentation or synthesis, such as handling of missing summary statistics, or data<br>conversions.   | 10-11          |  |  |
|                                  | 13c    | Describe any methods used to tabulate or visually display results of individual studies and syntheses.   | 10-11, Protoco |  |  |
|                                  | 13d    | Describe any methods used to synthesize results and provide a rationale for the choice(s). If meta-analysis was performed, describe the model(s), method(s) to identify the presence and extent of statistical heterogeneity, and software package(s) used.  | 10-11          |  |  |
|                                  | 13e    | Describe any methods used to explore possible causes of heterogeneity among study results.   | 10-11          |  |  |
|                                  | 13f    | Describe any sensitivity analyses conducted to assess robustness of the synthesized results.   | 11             |  |  |

| Section and Topic                                 | Item # | Checklist item   | Page #     |  |  |  |
|---|--------|--|------------|--|--|--|
| Reporting bias assessment                         | 14     | Describe any methods used to assess risk of bias due to missing results in a synthesis (arising from reporting biases).  | 10-11      |  |  |  |
| Certainty assessment                              | 15     | Describe any methods used to assess certainty (or confidence) in the body of evidence for an outcome.  |            |  |  |  |
| RESULTS   |        |  |            |  |  |  |
| Study selection                                   | 16a    | Describe the results of the search and selection process, from the number of records identified in the search to the number of studies included in the review, ideally using a flow diagram (see Figure 1).  | 12, Flow   |  |  |  |
|   | 16b    | Cite studies that met many but not all inclusion criteria ('near-misses') and explain why they were excluded.  | Flow diag  |  |  |  |
| Study characteristics                             | 17     | Cite each included study and present its characteristics.  | 12-13      |  |  |  |
| Risk of bias in studies                           | 18     | Present assessments of risk of bias for each included study.   | 12, Figure |  |  |  |
| Results of individual<br>studies                  | 19     | For all outcomes, present, for each study: (a) summary statistics for each group (where appropriate) and (b) an effect estimate and its precision (e.g. confidence/credible interval), ideally using structured tables or plots.   | 13-20, Ta  |  |  |  |
| Results of syntheses                              | 20a    | For each synthesis, briefly summarise the characteristics and risk of bias among contributing studies.   | 13-20, Ap  |  |  |  |
|   | 20b    | Present results of all statistical syntheses conducted. If meta-analysis was done, present for each the summary estimate and its precision (e.g. confidence/credible interval) and measures of statistical heterogeneity. If comparing groups, describe the direction of the effect. | 13-20, Ap  |  |  |  |
|   | 20c    | Present results of all investigations of possible causes of heterogeneity among study results.   | 13-20, Ap  |  |  |  |
|   | 20d    | Present results of all sensitivity analyses conducted to assess the robustness of the synthesized results.   | 13-20, Ap  |  |  |  |
| Reporting biases                                  | 21     | Present assessments of risk of bias due to missing results (arising from reporting biases) for each synthesis assessed.  | 12         |  |  |  |
| Certainty of evidence                             | 22     | Present assessments of certainty (or confidence) in the body of evidence for each outcome assessed.  | 13-20      |  |  |  |
| DISCUSSION  |        |  |            |  |  |  |
| Discussion  | 23a    | Provide a general interpretation of the results in the context of other evidence.  | 21-22      |  |  |  |
|   | 23b    | Discuss any limitations of the evidence included in the review.  | 22         |  |  |  |
|   | 23c    | Discuss any limitations of the review processes used.  | 22         |  |  |  |
|   | 23d    | Discuss implications of the results for practice, policy, and future research.   | 22-23      |  |  |  |
| OTHER INFORMATION                                 |        |  |            |  |  |  |
| Registration and protocol                         | 24a    | Provide registration information for the review, including register name and registration number, or state that the review was not registered.   | 6          |  |  |  |
|   | 24b    | Indicate where the review protocol can be accessed, or state that a protocol was not prepared.   | 6          |  |  |  |
|   | 24c    | Describe and explain any amendments to information provided at registration or in the protocol.  | 6, 9-10    |  |  |  |
| Support   | 25     | Describe sources of financial or non-financial support for the review, and the role of the funders or sponsors in the review.  |            |  |  |  |
| Competing interests                               | 26     | Declare any competing interests of review authors.   | 24-25      |  |  |  |
| Availability of data, code<br>and other materials | 27     | Report which of the following are publicly available and where they can be found: template data collection forms; data extracted from included studies; data used for all analyses; analytic code; any other materials used in the review.   |            |  |  |  |

PRISMA 2020 statement. MetaArXiv preprint (September 2020)

Appendix II. Search strategies for MEDLINE, Embase, CINAHL, and PsycINFO

# MEDLINE Ovid MEDLINE(R) ALL <1946 to Present>

# Searches

- 1 exp Back/
- 2 exp Back Injuries/
- 3 exp Back Muscles/
- 4 exp Back Pain/
- 5 Coccyx/
- 6 Discitis/
- 7 exp Intervertebral Disk/
- 8 Lumbar Vertebrae/
- 9 exp Lumbosacral Plexus/
- 10 Osteoarthritis/
- 11 Osteoarthritis, Spine/
- 12 Polyradiculopathy/
- 13 Radiculopathy/
- 14 Sacroiliac Joint/
- 15 Sacrum/
- 16 exp Sciatic Neuropathy/
- 17 exp Spinal Diseases/
- 18 Spine/
- 19 exp Spondylarthritis/
- 20 or/1-19
- 21 (back pain or backache or back ache or low\* back pain\*).ti,ab,kw,kf.
- 22 ((backache\* or back ache\*) adj3 (injur\* or pain\*)).ti,ab,kw,kf.
- 23 ((back or lumb\*) adj3 (ache\* or injur\* or pain\*)).ti,ab,kw,kf.
- 24 coccydynia.ti,ab,kw,kf.
  - ((coccyx or coccygeal or (L1 adj4 L5) or lumbarsacr\* or lumbosacr\* or lumbar or sacral or
- 25 sacrococcygeal or sacroiliac or sacro iliac or spinal or tailbone\* or vertebrogenic) adj3 (ache\* or break\* or broke\* or bruis\* or discomfort or fractur\* or hurt or injur\* or pain\* or tender\* or trauma)).ti,ab,kw,kf.
  - ((disc\* or disk\*) adj3 (avuls\* or bulg\* or compress\* or extrude\* or degenerat\* or displac\* or
- 26 herniat\* or hurt or injur\* or pain\* or protrud\* or prolaps\* or ruptur\* or sequester\* or slip\* or trauma or tear? or torn)).ti,ab,kw,kf.
- 27 (discitides or discitis or diskitides or diskitis or spondylodiscitides or spondylodiscitis or spondylodiskitis).ti,ab,kw,kf.
- 28 dorsalgia.ti,ab,kw,kf.
- 29 lumbago.ti,ab,kw,kf.
- 30 (lumbar adj (disc\* or disk\*) adj3 (avuls\* or extrude\* or degenerat\* or displac\* or herniat\* or prolaps\* or sequester\* or slip\* or injur\* or pain\*)).ti,ab,kw,kf.
- 31 (lumbar adj3 (compres\* or facet\* or nerve root\* or osteoarthritis or radicul\* or spinal stenos#s or spondylo\* or zygapophys\*)).ti,ab,kw,kf.
- 32 lumboischialgia.ti,ab,kw,kf.
- 33 Piriformis syndrome\*.ti,ab,kw,kf.
- 34 radiculalgia.ti,ab,kw,kf.
- 35 sciatic\*.ti,ab,kw,kf.
- 36 spondylosis.ti,ab,kw,kf.

- 37 ((stenos#s adj (spine or root or spinal)) or (failed adj3 "back syndrome\*")).ti,ab,kw,kf.
- 38 or/21-37
- 39 20 or 38
- 40 Adaptation, Psychological/
- 41 Affective Symptoms/
- 42 Anxiety/
- 43 Anxiety Disorders/
- 44 Depression/
- 45 Depressive Disorder/
- 46 Depressive Disorder, Major/
- 47 Dysthymic Disorder/
- 48 Irritable Mood/
- 49 Mental Health/
- 50 Mood Disorders/
- 51 Neurotic Disorders/
- 52 Sadness/
- 53 Stress, Psychological/
- 54 Suicidal Ideation/
- 55 Resilience, Psychological/
- 56 or/40-55
- 57 (depression or depressive or depressed).ti,kf,kw.
- 58 (depression or depressive or depressed).ab. /freq=2
- ((emotional\* or mental\* or psycholog\*) adj3 (adapt\* or affliction\* or cope? or coping or
   distress\* or health\* or illness\* or resilien\* or stress\* or unhealth\* or unwell or well being or
   wellbeing or wellness)).ti,ab,kw,kf.
  - (anguish\* or anxiety or anxious\* or dejected or desolat\* or despair\* or desperat\* or desponden\* or distress\* or dysthymi\* or hopeless\* or irritability or joyless\* or melanchol\* or mood\* or misery or miserable or morose or sadness or self-despair\* or self-pity\* or
- 60 unhap\*).ti,kw,kf. or (anguish\* or anxiety or anxious\* or dejected or desolat\* or despair\* or desperat\* or despenden\* or distress\* or dysthymi\* or hopeless\* or irritability or joyless\* or melanchol\* or mood\* or misery or miserable or morose or sadness or self-despair\* or self-pity\* or unhap\*).ab. /freq=2
- 61 (neuros#s or neurotic).ti,ab,kw,kf.
- 62 or/57-61
- 63 56 or 62
- 64 exp Case-Control Studies/
- 65 Control Groups/ or Matched-Pair Analysis/ or ((case\* adj3 control\*) or (case\* adj3 comparison\*) or control group\*).ti,ab,kw.
- 66 exp Cohort Studies/
- 67 ((cohort or longitudinal or prospective or retrospective) adj4 (study or studies or analy\*)).ti,ab.
- 68 or/64-67
- 69 39 and 63 and 68
- 70 (exp Animals/ not Humans/) or ((murin\* or mouse or mice or rat or rats or dog or dogs or cat or
- cats).ti. and ("in data review" or in process or publisher or "pubmed not medline").st.)
- 71 69 not 70

#### EMBASE

Ovid Embase <1974 to Present>

- # Searches
- 1 backache/
- 2 low back pain/
- 3 pelvic girdle pain/
- 4 discogenic pain/
- 5 cervical disk hernia/
- 6 diskitis/
- 7 exp experimental sciatic nerve injury/
- 8 failed back surgery syndrome/
- 9 exp intervertebral disk disease/
- 10 lumbar disk hernia/
- 11 lumbar plexus block/
- 12 lumbar spinal stenosis/
- 13 lumbar sympathectomy/
- 14 osteoarthritis/
- 15 exp radiculopathy/
- 16 sacroiliitis/
- 17 sciatic neuropathy/
- 18 exp sciatic nerve injury/
- 19 sciatica/
- 20 exp spine disease/
- 21 exp back/
- 22 exp pelvic girdle/
- 23 coccygeal vertebra/
- 24 coccygeus muscle/
- 25 coccyx/
- 26 exp intervertebral disk/
- 27 lumbar disk/
- 28 exp lumbar vertebra/
- 29 lumbar spine/
- 30 lumbosacral plexus/
- 31 lumbosacral spine/
- 32 sacroiliac joint/
- 33 sciatic nerve/
- 34 dislocation/
- 35 exp bone pain/
- 36 exp chronic pain/
- 37 exp inflammatory pain/
- 38 injury/
- 39 exp limb pain/
- 40 exp musculoskeletal pain/
- 41 pain/
- 42 phantom pain/
- 43 posttraumatic pain/
- 44 referred pain/
- 45 rupture/
- 46 spinal pain/
- 47 or/21-33

- 48 or/34-46
- 49 47 and 48
- 50 or/1-20
- 51 49 or 50
- 52 (back pain or backache or back ache or low\* back pain\*).ti,ab,kw.
- 53 ((backache\* or back ache\*) adj3 (injur\* or pain\*)).ti,ab,kw.
- 54 ((back or lumb\*) adj3 (ache\* or injur\* or pain\*)).ti,ab,kw.
- 55 coccydynia.ti,ab,kw.

((coccyx or coccygeal or (L1 adj4 L5) or lumbarsacr\* or lumbosacr\* or lumbar or sacral or sacrococcygeal or sacroiliac or sacro iliac or spinal or tailbone\* or vertebrogenic) adj3 (ache\*

- 56 satisfies of satisfies of satisfies of satisfies of spinal of tangone of vertebrogene) adj3 (acte or break\* or broke\* or bruis\* or discomfort or fractur\* or hurt or injur\* or pain\* or tender\* or trauma)).ti,ab,kw.
  - ((disc\* or disk\*) adj3 (avuls\* or bulg\* or compress\* or extrude\* or degenerat\* or displac\* or
- 57 herniat\* or hurt or injur\* or pain\* or protrud\* or prolaps\* or ruptur\* or sequester\* or slip\* or trauma or tear? or torn)).ti,ab,kw.
- 58 (discitides or discitis or diskitides or diskitis or spondylodiscitides or spondylodiscitis or spondylodiskitis).ti,ab,kw.
- 59 lumbago.ti,ab,kw.
- 60 (lumbar adj (disc\* or disk\*) adj3 (avuls\* or extrude\* or degenerat\* or displac\* or herniat\* or prolaps\* or sequester\* or slip\* or injur\* or pain\*)).ti,ab,kw.
- 61 (lumbar adj3 (compres\* or facet\* or nerve root\* or osteoarthritis or radicul\* or spinal stenos#s or spondylo\* or zygapophys\*)).ti,ab,kw.
- 62 lumboischialgia.ti,ab,kw.
- 63 Piriformis syndrome\*.ti,ab,kw.
- 64 radiculalgia.ti,ab,kw.
- 65 sciatic\*.ti,ab,kw.
- 66 spondylosis.ti,ab,kw.
- 67 ((stenos#s adj (spine or root or spinal)) or (failed adj3 "back syndrome\*")).ti,ab,kw.
- 68 or/52-67
- 69 51 or 68
- 70 affective neurosis/
- 71 anxiety/
- 72 anxiety disorder/
- 73 anxiety neurosis/
- 74 exp depression/
- 75 exp depression assessment/
- 76 distress syndrome/
- 77 emotional stress/
- 78 major affective disorder/
- 79 minor affective disorder/
- 80 mental health/
- 81 "mixed anxiety and depression"/
- 82 mood change/
- 83 mood disorder/
- 84 neurosis/
- 85 psychological resilience/
- 86 sadness/
- 87 suicidal ideation/
- 88 or/70-87
- 89 (depression or depressive or depressed).ti,kw.
- 90 (depression or depressive or depressed).ab. /freq=2

((emotional\* or mental\* or psycholog\*) adj3 (adapt\* or affliction\* or cope? or coping or

91 distress\* or health\* or illness\* or resilien\* or stress\* or unhealth\* or unwell or well being or wellbeing or wellness)).ti,ab,kw.

(anguish\* or anxiety or anxious\* or dejected or desolat\* or despair\* or desperat\* or desponden\* or distress\* or dysthymi\* or hopeless\* or irritability or joyless\* or melanchol\* or mood\* or misery or miserable or morose or sadness or self-despair\* or self-pity\* or

- 92 unhap\*).ti,kw. or (anguish\* or anxiety or anxious\* or dejected or desolat\* or despair\* or desperat\* or despenden\* or distress\* or dysthymi\* or hopeless\* or irritability or joyless\* or melanchol\* or mood\* or misery or miserable or morose or sadness or self-despair\* or self-pity\* or unhap\*).ab. /freq=2
- 93 (neuros#s or neurotic).ti,ab,kw.
- 94 or/89-93
- 95 88 or 94
- 96 69 and 95
- 97 exp case control study/
- 98 controlled study/ or pretest posttest control group design/ or static group comparison/ or ((case\* adj3 control\*) or (case\* adj3 comparison\*) or control group\*).ti,ab,kw.
- 99 cohort analysis/
- 100 ((cohort or longitudinal or prospective or retrospective) adj4 (study or studies or analy\*)).ti,ab.
- 101 or/97-100
- 102 96 and 101
- 103 (exp animal/ or animal experiment/ or nonhuman/) not exp human/
- 104 102 not 103

### CINAHL

EBSCOhost CINAHL Plus with Full Text

- # Query
- S1 (MH "Back")
- S2 (MH "Back Injuries+")
- S3 (MH "Back Pain+")
- S4 (MH "Coccyx")
- S5 (MH "Discitis")
- S6 (MH "Intervertebral Disk+")
- S7 (MH "Lumbar Vertebrae")
- S8 (MH "Lumbosacral Plexus+")
- S9 (MH "Osteoarthritis, Spine+")
- S10 (MH "Piriformis Syndrome")
- S11 (MH "Radiculopathy")
- S12 (MH "Polyradiculopathy")
- S13 (MH "Sacroiliac Joint")
- S14 (MH "Sacroiliac Joint Dysfunction")
- S15 (MH "Sacrum")
- S16 (MH "Sciatic Nerve+")
- S17 (MH "Sciatica")
- S18 (MH "Spinal Diseases+")
- S19 (MH "Spinal Injuries+")
- S20 (MH "Spine")
- S21
   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 OR S15 OR S16 OR S17 OR S18 OR S19 OR S20
- TI ( ("back pain" OR backache OR "back ache" OR "low\* back pain") ) OR AB ( ("back pain" S22 OR backache OR "back ache" OR "low\* back pain") ) OR SU ( ("back pain" OR backache OR

"back ache" OR "low\* back pain") ) TI ( ((backache\* OR "back ache\*") N3 (injur\* OR pain\*)) ) OR AB ( ((backache\* OR "back

- S23 ache\*") N3 (injur\* OR pain\*)) ) OR SU ( ((backache\* OR "back ache\*") N3 (injur\* OR pain\*)) )
- S24 TI ( ((back OR lumb\*) N3 (ache\* OR injur\* OR pain\*)) ) OR AB ( ((back OR lumb\*) N3
- (ache\* OR injur\* OR pain\*)) ) OR SU ( ((back OR lumb\*) N3 (ache\* OR injur\* OR pain\*)) )
   TI coccydynia OR AB coccydynia OR SU coccydynia
- TI ( ((coccyx OR coccygeal OR (L1 N4 L5) OR lumbarsacr\* OR lumbosacr\* OR lumbar OR sacral OR sacrococcygeal OR sacroiliac OR sacroiliac OR spinal OR tailbone\* OR vertebrogenic) N3 (ache\* OR break\* OR broke\* OR bruis\* OR discomfort OR fractur\* OR hurt OR injur\* OR pain\* OR tender\* OR trauma)) ) OR AB ( ((coccyx OR coccygeal OR (L1 N4 L5) OR lumbarsacr\* OR lumbosacr\* OR lumbar OR sacrococcygeal OR
- S26 sacroiliac OR sacro iliac OR spinal OR tailbone\* OR vertebrogenic) N3 (ache\* OR break\* OR broke\* OR bruis\* OR discomfort OR fractur\* OR hurt OR injur\* OR pain\* OR tender\* OR trauma)) ) OR SU ( ((coccyx OR coccygeal OR (L1 N4 L5) OR lumbarsacr\* OR lumbosacr\* OR lumbar OR sacral OR sacrococcygeal OR sacroiliac OR sacro iliac OR spinal OR tailbone\* OR vertebrogenic) N3 (ache\* OR break\* OR broke\* OR bruis\* OR discomfort OR fractur\* OR hurt OR injur\* OR pain\* OR tender\* OR hurt OR injur\* OR pain\* OR tender\* OR trauma)) )
   TI ( ((disc\* OR disk\*) N3 (avuls\* OR bulg\* OR compress\* OR extrude\* OR degenerat\* OR

TI ( ((disc\* OR disk\*) N3 (avuls\* OR bulg\* OR compress\* OR extrude\* OR degenerat\* OR displac\* OR herniat\* OR hurt OR injur\* OR pain\* OR protrud\* OR prolaps\* OR ruptur\* OR

S27 sequester\* OR slip\* OR trauma OR tear# OR torn)) ) OR AB ( ((disc\* OR disk\*) N3 (avuls\* OR bulg\* OR compress\* OR extrude\* OR degenerat\* OR displac\* OR herniat\* OR hurt OR injur\* OR pain\* OR protrud\* OR prolaps\* OR ruptur\* OR sequester\* OR slip\* OR trauma OR

tear# OR torn)) ) OR SU ( ((disc\* OR disk\*) N3 (avuls\* OR bulg\* OR compress\* OR extrude\* OR degenerat\* OR displac\* OR herniat\* OR hurt OR injur\* OR pain\* OR protrud\* OR prolaps\* OR ruptur\* OR sequester\* OR slip\* OR trauma OR tear# OR torn)) )

- S28 (discitides OR discitis OR diskitides OR diskitis OR spondylodiscitides OR spondylodiscitis
- OR spondylodiskitides OR spondylodiskitis)
- S29 TI dorsalgia OR AB dorsalgia OR SU dorsalgia
- S30 TI lumbago OR AB lumbago OR SU lumbago
   TI ( (lumbar N1 (disc\* OR disk\*) N3 (avuls\* OR extrude\* OR degenerat\* OR displac\* OR herniat\* OR prolaps\* OR sequester\* OR slip\* OR injur\* OR pain\*)) ) OR AB ( (lumbar N1 (disc\* OR disk\*) N3 (avuls\* OR extrude\* OR degenerat\* OR displac\* OR herniat\* OR
- S31 (disc\* OR disk\*) N3 (avuis\* OR extrude\* OR degenerat\* OR displac\* OR hermat\* OR prolaps\* OR sequester\* OR slip\* OR injur\* OR pain\*)) ) OR SU ( (lumbar N1 (disc\* OR disk\*) N3 (avuls\* OR extrude\* OR degenerat\* OR displac\* OR herniat\* OR prolaps\* OR sequester\* OR slip\* OR injur\* OR pain\*)) )

TI ( (lumbar N3 (compres\* OR facet\* OR nerve root\* OR osteoarthritis OR radicul\* OR spinal stenos?s OR spondylo\* OR zygapophys\*)) ) OR AB ( (lumbar N3 (compres\* OR facet\* OR

- S32 nerve root\* OR osteoarthritis OR radicul\* OR spinal stenos?s OR spondylo\* OR zygapophys\*)) ) OR SU ( (lumbar N3 (compres\* OR facet\* OR nerve root\* OR osteoarthritis OR radicul\* OR spinal stenos?s OR spondylo\* OR zygapophys\*)) )
- S33 TI lumboischialgia OR AB lumboischialgia OR SU lumboischialgia
- S34 TI "Piriformis syndrome\*" OR AB "Piriformis syndrome\*" OR SU "Piriformis syndrome\*"
- S35 TI radiculalgia OR AB radiculalgia OR SU radiculalgia
- S36 TI sciatic\* OR AB sciatic\* OR SU sciatic\*
- S37 TI spondylosis OR AB spondylosis OR SU spondylosis
   TI ( ((stenos?s N1 (spine OR root OR spinal)) OR (failed N3 "back syndrome\*")) ) OR AB (
- S38 ((stenos?s N1 (spine OR root OR spinal)) OR (failed N3 "back syndrome\*")) ) OR SU ( ((stenos?s N1 (spine OR root OR spinal)) OR (failed N3 "back syndrome\*")) )
- S22 OR S23 OR S24 OR S25 OR S26 OR S27 OR S28 OR S29 OR S30 OR S31 OR S32 OR S29 OR S30 OR S31 OR S32 OR S3
- S33 OR S34 OR S35 OR S36 OR S37 OR S38
- S40 S21 OR S39
- S41 (MH "Adaptation, Psychological")
- S42 (MH "Adjustment Disorders")
- S43 (MH "Affective Disorders")
- S44 (MH "Affective Symptoms")
- S45 (MH "Anxiety")
- S46 (MH "Anxiety Disorders")
- S47 (MH "Depression")
- S48 (MH "Depression, Reactive")
- S49 (MH "Dysthymic Disorder")
- S50 (MH "Hardiness")
- S51 (MH "Mental Health")
- S52 (MH "Neurotic Disorders")
- S53 (MH "Sadness")
- S54 (MH "Stress, Psychological")
- S55 (MH "Suicidal Ideation")
- S56
   S41 OR S42 OR S43 OR S44 OR S45 OR S46 OR S47 OR S48 OR S49 OR S50 OR S51 OR

   S52 OR S53 OR S54 OR S55
- S57 TI ( (depression OR depressive OR depressed) ) OR SU ( (depression OR depressive OR depressed))
- S58 AB (depression OR depressive OR depressed)
- S59 S58 OR S57

TI ( ((emotional\* OR mental\* OR psycholog\*) N3 (adapt\* OR affliction\* OR cope# OR coping OR distress\* OR health\* OR illness\* OR resilien\* OR stress\* OR unhealth\* OR unwell OR "well being" OR wellbeing OR wellness)) ) OR AB ( ((emotional\* OR mental\* OR psycholog\*) N3 (adapt\* OR affliction\* OR cope# OR coping OR distress\* OR health\* OR

- S60 psycholog<sup>1</sup>) NS (adapt<sup>2</sup> OK affiltion<sup>2</sup> OK cope# OK coping OK distress<sup>2</sup> OK health<sup>2</sup> OK isliess<sup>2</sup> OK health<sup>2</sup> OK isliess<sup>3</sup> OK health<sup>2</sup> OK isliess<sup>3</sup> OK health<sup>3</sup> OK wellbeing OR wellbess)) ) OR SU ( ((emotional\* OR mental\* OR psycholog\*) N3 (adapt\* OR affliction\* OR cope# OR coping OR distress\* OR health\* OR illness\* OR resilien\* OR stress\* OR unhealth\* OR unwell OR "well being" OR wellbeing OR wellbeing OR wellness)) )
   TI ( (anguish\* OR anxiety OR anxious\* OR dejected OR desolat\* OR despair\* OR desperat\* OR desponden\* OR distress\* OR dysthymi\* OR hopeless\* OR irritability OR joyless\* OR melanchol\* OR mood\* OR misery OR miserable OR morose OR sadness OR self-despair\* OR
- self-pity\* OR unhap\*) ) OR SU ( (anguish\* OR anxiety OR anxious\* OR dejected OR desolat\* OR despair\* OR desperat\* OR desponden\* OR distress\* OR dysthymi\* OR hopeless\* OR irritability OR joyless\* OR melanchol\* OR mood\* OR misery OR miserable OR morose OR sadness OR self-despair\* OR self-pity\* OR unhap\*) )
   AB (anguish\* OR anxiety OR anxious\* OR dejected OR desolat\* OR despair\* OR desperat\*
- S62 OR desponden\* OR distress\* OR dysthymi\* OR hopeless\* OR irritability OR joyless\* OR melanchol\* OR mood\* OR misery OR miserable OR morose OR sadness OR self-despair\* OR self-pity\* OR unhap\*)
- S63 S61 OR S62
- S64 TI ( (neuros?s OR neurotic) ) OR AB ( (neuros?s OR neurotic) ) OR SU ( (neuros?s OR neurotic) )
- S65 S59 OR S60 OR S63 OR S64
- S66 S56 OR S65
- S67 (MH "Case Control Studies")
- S68 (MH "Hospital-Based Case Control")
- S69 (MH "Matched Case Control")
- S70 (MH "Population-Based Case Control")
- S71 (MH "Prospective Studies+")
- S72 (MH "Panel Studies+")
- S73 (MH "Pseudolongitudinal Studies")
- S74 (MH "Retrospective Design")
- S75 TI ( ((case\* N3 control\*) OR (case\* N3 comparison\*) OR control group\*) ) OR SU ( ((case\* N3 control\*) OR (case\* N3 comparison\*) OR control group\*) )
- S76 S67 OR S68 OR S69 OR S70 OR S71 OR S72 OR S73 OR S74 OR S75
- S77 S40 AND S66 AND S76
- S78 S77 NOT ((MH "Animals+") NOT (MH "Human"))

### PSYCINFO

Ovid PsycINFO <1806 to Present>

#### # Searches

- 1 "back (anatomy)"/
- 2 back pain/
- 3 lumbar spinal cord/
- 4 musculoskeletal system/
- 5 neuropathic pain/
- 6 spinal column/
- 7 spinal nerves/
- 8 or/1-7
- 9 (back pain or backache or back ache or low\* back pain\*).ti,ab,id.
- 10 ((backache\* or back ache\*) adj3 (injur\* or pain\*)).ti,ab,id.
- 11 ((back or lumb\*) adj3 (ache\* or injur\* or pain\*)).ti,ab,id.
- 12 coccydynia.ti,ab,id.
  - ((coccyx or coccygeal or (L1 adj4 L5) or lumbarsacr\* or lumbosacr\* or lumbar or sacral or
- 13 sacrococcygeal or sacroiliac or sacro iliac or spinal or tailbone\* or vertebrogenic) adj3 (ache\* or break\* or broke\* or bruis\* or discomfort or fractur\* or hurt or injur\* or pain\* or tender\* or trauma)).ti,ab,id.

((disc\* or disk\*) adj3 (avuls\* or bulg\* or compress\* or extrude\* or degenerat\* or displac\* or

- 14 herniat\* or hurt or injur\* or pain\* or protrud\* or prolaps\* or ruptur\* or sequester\* or slip\* or trauma or tear? or torn)).ti,ab,id.
- 15 (discitides or diskitides or diskitis or spondylodiscitides or spondylodiscitis or
- spondylodiskitides or spondylodiskitis).ti,ab,id.
- 16 dorsalgia.ti,ab,id.
- 17 lumbago.ti,ab,id.
- 18 (lumbar adj (disc\* or disk\*) adj3 (avuls\* or extrude\* or degenerat\* or displac\* or herniat\* or prolaps\* or sequester\* or slip\* or injur\* or pain\*)).ti,ab,id.
- 19 (lumbar adj3 (compres\* or facet\* or nerve root\* or osteoarthritis or radicul\* or spinal stenos#s or spondylo\* or zygapophys\*)).ti,ab,id.
- 20 lumboischialgia.ti,ab,id.
- 21 Piriformis syndrome\*.ti,ab,id.
- 22 radiculalgia.ti,ab,id.
- 23 sciatic\*.ti,ab,id.
- 24 spondylosis.ti,ab,id.
- 25 ((stenos#s adj (spine or root or spinal)) or (failed adj3 "back syndrome\*")).ti,ab,id.
- 26 or/9-25
- 27 8 or 26
- 28 adjustment/
- 29 adjustment disorders/
- 30 anxiety/
- 31 anxiety disorders/
- 32 "depression (emotion)"/
- 33 dysthymic disorder/
- 34 emotional adjustment/
- 35 emotional states/
- 36 irritability/
- 37 major depression/
- 38 mental health/

- 39 neurosis/
- 40 psychological endurance/
- 41 reactive depression/
- 42 "resilience (psychological)"/
- 43 sadness/
- 44 stress reactions/
- 45 suicidal ideation/
- 46 or/28-45
- (depression or depressive or depressed).ti,ab,id.
   ((emotional\* or mental\* or psycholog\*) adj3 (adapt\* or affliction\* or cope? or coping or distress\* or health\* or illness\* or resilien\* or stress\* or unhealth\* or unwell or well being or
- 48 wellbeing or wellness)).ti,id. or ((emotional\* or mental\* or psycholog\*) adj3 (adapt\* or affliction\* or cope? or coping or distress\* or health\* or illness\* or resilien\* or stress\* or unhealth\* or unwell or well being or wellbeing or wellness)).ab. /freq=2 (anguish\* or anxiety or anxious\* or dejected or desolat\* or despair\* or desperat\* or desponden\* or distress\* or dysthymi\* or hopeless\* or irritability or joyless\* or melanchol\* or mood\* or misery or miserable or morose or sadness or self-despair\* or self-pity\* or unhap\*).ti,id. or
- 49 Inserv of miserable of morose of sadness of self-despair\* of self-pity\* of unhap\*).u,id. of (anguish\* or anxiety or anxious\* or dejected or desolat\* or despair\* or desperat\* or despenden\* or distress\* or dysthymi\* or hopeless\* or irritability or joyless\* or melanchol\* or mood\* or misery or miserable or morose or sadness or self-despair\* or self-pity\* or unhap\*).ab. /freq=2
- 50 (neuros#s or neurotic).ti,ab,id.
- 51 or/47-50
- 52 46 or 51
- 53 27 and 52
- 54 case control stud\*.mp.
- 55 experiment controls/
- 56 longitudinal studies/ or prospective studies/ or retrospective studies/
- 57 ((case\* adj3 control\*) or (case\* adj3 comparison\*) or control group\*).ti,ab,id.
- 58 ((cohort or longitudinal or prospective or retrospective) adj4 (study or studies or analy\*)).ti,ab.
- 59 cohort analysis/
- 60 or/54-59
- 61 53 and 60
- 62 limit 61 to human

**Appendix IIIa.** Table outlining population, exposure, outcomes, and eligible study designs for the systematic review assessing the association between depressive symptoms or depression on health outcomes in persons with low back pain

| Population ]  | <b>Description</b><br>We targeted studies with an inception cohort of individuals aged 16 years and older with LBP with or without radiculopathy. LBP is defined as pain localized between the costal margin and inferior gluteal folds, with or without referred leg pain in the absence of serious or major pathology. Radiculopathy is inflammation, injury, or compression of  |
|---|--|
|   | spinal nerve roots with pain, weakness, or numbress in a myotomal or dermatomal  |
|   | distribution, and may be attributed to spinal stenosis or lumbar disc herniation. We   |
|   | excluded LBP due to major structural or serious pathology (e.g., spinal fractures, spinal dislocations, spinal cord injury, inflammatory arthritis, neoplasms or malignancies) and surgical populations. We included studies involving participants under 16 years of age, as long as those studies reported stratified results for individuals aged 16 years and older.   |
| Exposure  | We included studies that assessed depressive symptoms or depression as the exposure or prognostic factor. Depressive symptoms are defined as self-reported characteristic attitudes and symptoms of depression, such as taking little interest or pleasure in doing things or feeling down, depressed, or hopeless. We included depressive symptoms measured on standardized questionnaires <sup>a</sup> (e.g., Center for Epidemiologic Studies Depression Scale, Beck Depression Index). We also included diagnosed depression, including diagnoses captured using standardized questionnaires, which has two main categories: 1) major depressive disorder/episode; and 2) dysthymia. Major depressive disorder/episode presents with symptoms such as depressed mood, loss of interest and enjoyment, and decreased energy, and can be categorized as mild, moderate, or severe based on the symptom frequency and severity. Dysthymia is a persistent or chronic form of mild depression with symptoms similar to depressive episodes, but are less intense and persist longer.   |
| )<br> <br> | We targeted the following health outcomes: 1) pain (e.g., pain intensity); 2) disability<br>(e.g., impairment, activity limitations, participation restriction); 3) overall health status<br>(e.g., health-related quality of life, recovery); 4) satisfaction with care; and 5) health care<br>utilization (e.g., physician visits, emergency department visits, hospitalizations, spinal<br>imaging). These were informed by core outcome domains that are considered important<br>for LBP research among international panels of experts. Only standardized outcome<br>measures (e.g., standardized questionnaires or administrative data) were included. We<br>evaluated the validity and reliability of the standardized questionnaires during the risk of<br>bias assessment. Questionnaires for health outcomes included: 1) Visual Analogue Scale<br>and Numeric Rating Scale for measuring pain intensity; 2) Roland Morris Disability<br>Questionnaire and Oswestry Disability Index for measuring disability; 3) 36-item Short<br>Form Survey (SF-36), 12-item Short Form Survey (SF-12), and Global Perceived<br>Recovery for measuring overall health status; 4) Patient Satisfaction Questionnaire for<br>measuring satisfaction with care; and 5) National Ambulatory Medical Care Survey and<br>73-item LBP health care utilization questionnaire for measuring health care utilization. |
| Study 0   | Cohort studies, case-control studies, or secondary analyses of data from randomized trials   |
| Designs<br>LBP – low bac  | le noin  |

LBP – low back pain

<sup>a</sup>Common self-reported questionnaires for depressive symptoms have a recall period ranging from past one week (Center for Epidemiologic Studies Depression Scale, Hospital Anxiety and Depression Scale) to past two weeks (Patient Health Questionnaire-2, Patient Health Questionnaire-9) Appendix IIIb. Categories to guide the assessment of homogeneity across studies

| Category | Description*  |
|----------|---|
| 1A       | Population: type of LBP   |
|          | - LBP OR  |
|          | - LBP with radiculopathy  |
| 1B       | Population: duration of LBP   |
|          | - Acute/subacute (<12 weeks' duration) OR   |
|          | - Chronic (≥12 weeks' duration)   |
| 2A       | Exposure: type of condition   |
|          | - Depressive symptoms OR  |
|          | - Depression  |
| 2B       | Exposure: severity of condition   |
|          | - Mild (e.g., mild depression) OR   |
|          | - Severe (e.g., severe depression)  |
| 3        | Outcome: type   |
|          | - Pain intensity  |
|          | - LBP-related disability  |
|          | - Health-related quality of life  |
|          | - Type of health care utilization (e.g., family physician visit, specialist visit, or |
|          | spinal radiograph)  |

LBP – low back pain

\*Describes how studies within the listed categories would be considered homogeneous (e.g., studies targeting the following would be considered homogeneous: 1) chronic LBP without radiculopathy as the population; 2) severe depression as the exposure; 3) pain intensity as the outcome)

Appendix IV. List of all included papers in systematic review (62 studies reported in 66 articles)

- 1. Lehmann TR, Spratt KF, Lehmann KK. Predicting long-term disability in low back injured workers presenting to a spine consultant. Spine (Phila Pa 1976) 1993;18(8):1103-12.
- 2. Rainville J, Ahern DK, Phalen L. Altering beliefs about pain and impairment in a functionally oriented treatment program for chronic low back pain. Clin J Pain 1993;9(3):196-201.
- 3. Gatchel RJ, Polatin PB, Mayer TG. The dominant role of psychosocial risk factors in the development of chronic low back pain disability. Spine (Phila Pa 1976) 1995;20(24):2702-9.
- 4. Cherkin DC, Deyo RA, Street JH, et al. Predicting poor outcomes for back pain seen in primary care using patients' own criteria. Spine (Phila Pa 1976) 1996;21(24):2900-7.
- 5. Harkapaa KJ, A; Estlander, A. Health optimism and control beliefs as predictors for treatment outcome of a multimodal back treatment program. Psychology and Health 1996;12:123-34.
- 6. Dionne CE, Koepsell TD, Von Korff M, et al. Predicting long-term functional limitations among back pain patients in primary care settings. J Clin Epidemiol 1997;50(1):31-43.
- 7. Epping-Jordan JE, Wahlgren DR, Williams RA, et al. Transition to chronic pain in men with low back pain: predictive relationships among pain intensity, disability, and depressive symptoms. Health Psychol 1998;17(5):421-7.
- 8. Vendrig AA. Prognostic factors and treatment-related changes associated with return to work in the multimodal treatment of chronic back pain. J Behav Med 1999;22(3):217-32.
- 9. Fritz JM, George SZ. Identifying psychosocial variables in patients with acute work-related low back pain: the importance of fear-avoidance beliefs. Phys Ther 2002;82(10):973-83.
- 10. Fransen M, Woodward M, Norton R, et al. Risk factors associated with the transition from acute to chronic occupational back pain. Spine (Phila Pa 1976) 2002;27(1):92-8.
- 11. Cassidy JD, Carroll L, Côté P, et al. Low back pain after traffic collisions: a population-based cohort study. Spine (Phila Pa 1976) 2003;28(10):1002-9.
- 12. Michaelson P, Sjölander P, Johansson H. Factors predicting pain reduction in chronic back and neck pain after multimodal treatment. Clin J Pain 2004;20(6):447-54.
- 13. Tubach F, Beauté J, Leclerc A. Natural history and prognostic indicators of sciatica. J Clin Epidemiol 2004;57(2):174-9.
- 14. Sieben JM, Vlaeyen JW, Portegijs PJ, et al. A longitudinal study on the predictive validity of the fearavoidance model in low back pain. Pain 2005;117(1-2):162-70.
- 15. Truchon M, Côté D. Predictive validity of the Chronic Pain Coping Inventory in subacute low back pain. Pain 2005;116(3):205-12.
- 16. Campello MA, Weiser SR, Nordin M, et al. Work retention and nonspecific low back pain. Spine (Phila Pa 1976) 2006;31(16):1850-7.
- 17. Grotle M, Vøllestad NK, Brox JI. Screening for yellow flags in first-time acute low back pain: reliability and validity of a Norwegian version of the Acute Low Back Pain Screening Questionnaire. Clin J Pain 2006;22(5):458-67.
- 18. Patel SM. Psychosocial predictors of pain chronicity in Navy servicemen: Alliant International University, San Diego; 2006.
- 19. Ritzwoller DP, Crounse L, Shetterly S, et al. The association of comorbidities, utilization and costs for patients identified with low back pain. BMC Musculoskelet Disord 2006;7:72.
- 20. Newell D, Field J. Who will get better? Predicting clinical outcomes in a chiropractic practice. Clinical Chiropractic 2007;10(4):179-86.
- 21. Weidenhammer W, Linde K, Streng A, et al. Acupuncture for chronic low back pain in routine care: a multicenter observational study. Clin J Pain 2007;23(2):128-35.

- 22. Henschke N, Maher CG, Refshauge KM, et al. Prognosis in patients with recent onset low back pain in Australian primary care: inception cohort study. Bmj 2008;337(7662):a171.
- 23. van der Hulst M, Vollenbroek-Hutten MM, Groothuis-Oudshoorn KG, et al. Multidisciplinary rehabilitation treatment of patients with chronic low back pain: a prognostic model for its outcome. Clin J Pain 2008;24(5):421-30.
- 24. Gurcay E, Bal A, Eksioglu E, et al. Acute low back pain: clinical course and prognostic factors. Disabil Rehabil 2009;31(10):840-5.
- 25. Reme SE, Hagen EM, Eriksen HR. Expectations, perceptions, and physiotherapy predict prolonged sick leave in subacute low back pain. BMC Musculoskelet Disord 2009;10:139.
- 26. Smeets RJ, Maher CG, Nicholas MK, et al. Do psychological characteristics predict response to exercise and advice for subacute low back pain? Arthritis Rheum 2009;61(9):1202-9.
- 27. Grotle M, Foster NE, Dunn KM, et al. Are prognostic indicators for poor outcome different for acute and chronic low back pain consulters in primary care? Pain 2010;151(3):790-7.
- 28. Shaw WS, Means-Christensen AJ, Slater MA, et al. Psychiatric disorders and risk of transition to chronicity in men with first onset low back pain. Pain Med 2010;11(9):1391-400.
- 29. Streitberger K, Müller T, Eichenberger U, et al. Factors determining the success of radiofrequency denervation in lumbar facet joint pain: a prospective study. Eur Spine J 2011;20(12):2160-5.
- Hicks GE, Benvenuti F, Fiaschi V, et al. Adherence to a community-based exercise program is a strong predictor of improved back pain status in older adults: an observational study. Clin J Pain 2012;28(3):195-203.
- Hiebert R, Campello MA, Weiser S, et al. Predictors of short-term work-related disability among active duty US Navy personnel: a cohort study in patients with acute and subacute low back pain. Spine J 2012;12(9):806-16.
- 32. Melloh M, Elfering A, Käser A, et al. Depression impacts the course of recovery in patients with acute low-back pain. Behav Med 2013;39(3):80-9.
- 33. Elfering A, Käser A, Melloh M. Relationship between depressive symptoms and acute low back pain at first medical consultation, three and six weeks of primary care. Psychol Health Med 2014;19(2):235-46.
- 34. Melloh M, Elfering A, Chapple CM, et al. Prognostic occupational factors for persistent low back pain in primary care. Int Arch Occup Environ Health 2013;86(3):261-9.
- 35. Melloh M, Elfering A, Salathé CR, et al. Predictors of sickness absence in patients with a new episode of low back pain in primary care. Ind Health 2012;50(4):288-98.
- 36. Melloh M, Elfering A, Stanton TR, et al. Who is likely to develop persistent low back pain? A longitudinal analysis of prognostic occupational factors. Work 2013;46(3):297-311.
- 37. Scheele J, Enthoven WT, Bierma-Zeinstra SM, et al. Course and prognosis of older back pain patients in general practice: a prospective cohort study. Pain 2013;154(6):951-7.
- 38. Fischer CA, Neubauer E, Adams HS, et al. Effects of multidisciplinary pain treatment can be predicted without elaborate questionnaires. Int Orthop 2014;38(3):617-26.
- 39. van Hooff ML, Spruit M, O'Dowd JK, et al. Predictive factors for successful clinical outcome 1 year after an intensive combined physical and psychological programme for chronic low back pain. Eur Spine J 2014;23(1):102-12.
- 40. Scherrer JF, Salas J, Lustman PJ, et al. Change in opioid dose and change in depression in a longitudinal primary care patient cohort. Pain 2015;156(2):348-55.
- 41. Viniol A, Jegan N, Brugger M, et al. Even Worse Risk Factors and Protective Factors for Transition from Chronic Localized Low Back Pain to Chronic Widespread Pain in General Practice: A Cohort Study. Spine (Phila Pa 1976) 2015;40(15):E890-9.

- 42. Cougot B, Petit A, Paget C, et al. Chronic low back pain among French healthcare workers and prognostic factors of return to work (RTW): a non-randomized controlled trial. J Occup Med Toxicol 2015;10:40.
- 43. Kerr D, Zhao W, Lurie JD. What Are Long-term Predictors of Outcomes for Lumbar Disc Herniation? A Randomized and Observational Study. Clin Orthop Relat Res 2015;473(6):1920-30.
- 44. Lubelski D, Thompson NR, Agrawal B, et al. Prediction of quality of life improvements in patients with lumbar stenosis following use of membrane stabilizing agents. Clin Neurol Neurosurg 2015;139:234-40.
- 45. Enthoven WT, Koes BW, Bierma-Zeinstra SM, et al. Defining trajectories in older adults with back pain presenting in general practice. Age Ageing 2016;45(6):878-83.
- 46. Steenstra IA, Franche RL, Furlan AD, et al. The Added Value of Collecting Information on Pain Experience When Predicting Time on Benefits for Injured Workers with Back Pain. J Occup Rehabil 2016;26(2):117-24.
- Traeger AC, Hübscher M, Henschke N, et al. Emotional distress drives health services overuse in patients with acute low back pain: a longitudinal observational study. Eur Spine J 2016;25(9):2767-73.
- 48. Yarlas A, Miller K, Wen W, et al. A Subgroup Analysis Found no Diminished Response to Buprenorphine Transdermal System Treatment for Chronic Low Back Pain Patients Classified with Depression. Pain Pract 2016;16(4):473-85.
- 49. Adnan R, Van Oosterwijck J, Cagnie B, et al. Determining Predictive Outcome Factors for a Multimodal Treatment Program in Low Back Pain Patients: A Retrospective Cohort Study. J Manipulative Physiol Ther 2017;40(9):659-67.
- 50. Friedman BW, Gensler S, Yoon A, et al. Predicting three-month functional outcomes after an ED visit for acute low back pain. Am J Emerg Med 2017;35(2):299-305.
- 51. Jegan NR, Brugger M, Viniol A, et al. Psychological risk and protective factors for disability in chronic low back pain a longitudinal analysis in primary care. BMC Musculoskelet Disord 2017;18(1):114.
- 52. Nordeman L, Thorselius L, Gunnarsson R, et al. Predictors for future activity limitation in women with chronic low back pain consulting primary care: a 2-year prospective longitudinal cohort study. BMJ Open 2017;7(6):e013974.
- 53. Nordstoga AL, Nilsen TIL, Vasseljen O, et al. The influence of multisite pain and psychological comorbidity on prognosis of chronic low back pain: longitudinal data from the Norwegian HUNT Study. BMJ Open 2017;7(5):e015312.
- Dengler J, Sturesson B, Kools D, et al. Risk Factors for Continued Opioid Use in Conservative Versus Surgical Management of Low Back Pain Originating From the Sacroiliac Joint. Global Spine J 2018;8(5):453-9.
- 55. Oliveira DS, Vélia Ferreira Mendonça L, Sofia Monteiro Sampaio R, et al. The Impact of Anxiety and Depression on the Outcomes of Chronic Low Back Pain Multidisciplinary Pain Management-A Multicenter Prospective Cohort Study in Pain Clinics with One-Year Follow-up. Pain Med 2019;20(4):736-46.
- 56. Pagé MG, Boyd K, Ware MA. Examination of the Course of Low Back Pain Intensity Based on Baseline Predictors and Health Care Utilization Among Patients Treated in Multidisciplinary Pain Clinics: A Quebec Pain Registry Study. Pain Med 2019;20(3):564-73.
- 57. Bahar-Ozdemir Y, Sencan S, Ercalik T, et al. The Effect of Pre-Treatment Depression, Anxiety and Somatization Levels on Transforaminal Epidural Steroid Injection: A Prospective Observational Study. Pain Physician 2020;23(3):E273-e80.

- 58. Hartvigsen L, Kongsted A, Vach W, et al. Baseline Characteristics May Help Indicate the Best Choice of Health Care Provider for Back Pain Patients in Primary Care: Results From a Prospective Cohort Study. J Manipulative Physiol Ther 2020;43(1):13-23.
- 59. Imagama S, Murakami H, Kaito T, et al. Impact of background factors on outcomes of pharmacological therapy for chronic low back pain: A nationwide multicenter prospective study. J Orthop Sci 2020.
- 60. Klyne DM, Hodges PW. Circulating Adipokines in Predicting the Transition from Acute to Persistent Low Back Pain. Pain Med 2020.
- 61. Ranger TA, Cicuttini FM, Jensen TS, et al. Catastrophization, fear of movement, anxiety, and depression are associated with persistent, severe low back pain and disability. Spine J 2020;20(6):857-65.
- 62. Zackova M, Aspide R, Braghittoni A, et al. Yellow flag on prognostic factors for non-specific chronic low back pain patients subjected to mini-invasive treatment: a cohort study. Eur Spine J 2020;29(8):1879-86.
- 63. Glattacker M, Heyduck K, Jakob T. Yellow flags as predictors of rehabilitation outcome in chronic low back pain. Rehabil Psychol 2018;63(3):408-17.
- 64. Halonen JI, Shiri R, Magnusson Hanson LL, et al. Risk and Prognostic Factors of Low Back Pain: Repeated Population-based Cohort Study in Sweden. Spine (Phila Pa 1976) 2019;44(17):1248-55.
- 65. Demarchi SJ, Oliveira CB, Franco MR, et al. Association of perceived physical overload at work with pain and disability in patients with chronic non-specific low back pain: a 6-month longitudinal study. Eur Spine J 2019;28(7):1586-93.
- 66. Kim EJ, Chotai S, Stonko DP, et al. Patient-reported outcomes after lumbar epidural steroid injection for degenerative spine disease in depressed versus non-depressed patients. Spine J 2017;17(4):511-7.

Appendix V. Reported associations between depressive symptoms or depression and health outcomes for included studies

| Study                               | Sample<br>Size | Prognostic Factor Measure   | Outcome Measure  | Key Results  |
|-------------------------------------|----------------|---|--|--|
| Acute LBP (<3                       |                |   |  |  |
| months' duration)                   |                |   |  |  |
| Lehmann, 1993 <sup>1</sup>          | 70             | Depressive symptoms:<br>Beck Depression Inventory (0-<br>63, higher=worse)  | Work status (return to work)   | At 6 months: r=0.17, p<0.26<br>Depressive symptoms did not correlate with return to work status.   |
| Gatchel, 1995 <sup>2</sup>          | 421            | Depressive symptoms:<br>Minnesota Multiphasic<br>Personality Inventory MMPI-D<br>(continuous score)                             | Work status (return to work)   | At 1 year:<br>Depressive symptoms were not associated with return to work<br>(depressive symptoms not retained in adjusted analysis; model estimates<br>not reported).   |
| Epping-Jordan,<br>1998 <sup>3</sup> | 78             | Depressive symptoms:<br>Beck Depression Inventory (0-<br>63, higher=worse)  | Pain (Descriptor<br>Differential Scale)<br>Disability (Sickness Impact<br>Profile) | At 6 months:Pain: Adjusted $\beta$ 0.06 p=0.146Disability: Adjusted $\beta$ 0.07 p=0.453At 12 months:Pain: Adjusted $\beta$ -0.06 p=0.584Disability: Adjusted $\beta$ 0.05 p=0.686Depressive symptoms were not associated with pain or disability.   |
| Fritz, 2001 <sup>4</sup>            | 78             | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression (0-60;<br>higher=worse)                                | Restrictions in work status<br>(versus return to work<br>without restrictions)     | Mean difference in baseline depressive symptoms between having<br>persistent restrictions versus no restrictions in work status at 4 weeks:<br>4.4 (95% CI 0.70, 8.0) p=0.02<br>Depressive symptoms significantly differed between those with<br>persistent restrictions versus no restrictions in work status |
| Cassidy, 2003 <sup>5</sup>          | 3232           | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression (0-60,<br>higher=worse; >16 as depressive<br>symptoms) | Time to motor vehicle<br>collision claim closure                                   | No fault claimantsUnadjusted HR 0.52 (95% CI 0.43, 0.63)Adjusted HR 0.62 (95% CI 0.50, 0.76)Tort claimantsUnadjusted HR 0.54 (95% CI 0.43, 0.68)Adjusted HR 0.70 (95% CI 0.53, 0.93)Depressive symptoms were associated with slower claim closure from<br>motor vehicle collision.                             |
| Sieben, 2005 <sup>6</sup>           | 222            | Depressive symptoms:<br>Beck Depression Inventory<br>("negative view of self" subscale)   | Pain and disability (Graded<br>Chronic Pain Scale)                                 | At 6 months: $\rho$ =.17, p<0.05<br>At 12 months: $\rho$ =.20, p<0.05<br>End of study: $\rho$ =.27, p<0.05<br>Depressive symptoms were associated with pain/disability.  |
| Truchon, 2005 <sup>7</sup>          | 321            | Depressive symptoms:<br>Hospital Anxiety and Depression<br>Scale-Depression (0-21;<br>higher=worse)                             | Work status (returned to work or not)  | At 6 months:<br>Unadjusted OR 0.74 (95% CI 0.62, 0.89), p<0.01<br>Depressive symptoms were associated with non-return to work at 6<br>months.  |

| Study                          | Sample<br>Size | Prognostic Factor Measure   | Outcome Measure   | Key Results  |
|--------------------------------|----------------|---|---|--|
| Grotle, 2006 <sup>8</sup>      | 123            | Depressive symptoms:<br>Acute Low Back Pain Screening<br>Questionnaire – depression<br>subscale                   | Pain (Numeric Rating<br>Scale), disability (Roland<br>Morris Disability<br>Questionnaire)                           | <u>At 12 months:</u><br>Pain: Adjusted OR 1.4 (95% CI 1.09, 1.67)<br>Disability: Adjusted OR 1.4 (95% CI 1.13, 1.71)<br>Depressive symptoms were associated with pain and disability.  |
| Patel, 2006 <sup>9</sup>       | 107            | Depressive symptoms:<br>Hamilton Rating Scale for<br>Depression (0-52, higher=worse)                              | Pain (Descriptor<br>Differential Scale),<br>disability (Sickness Impact<br>Profile)                                 | Pain: r=0.41, p<0.01<br>Disability: r=0.52, p<0.01<br>Chronicity (both pain and disability): r=0.43, p<0.01<br>Depressive symptoms were associated with pain and disability.   |
| Ritzwoller, 2006 <sup>10</sup> | 16,567         | Depression (ICD-9 codes)  | Health care utilization<br>(inpatient admissions)   | All inpatient admissions over 24 months<br>Adjusted OR 1.27 (95% CI 1.13, 1.46)<br><u>MSK-related inpatient admissions over 24 months</u><br>Adjusted OR 0.96 (95% CI 0.74, 1.27)<br>Depression was associated with all inpatient admissions but not<br>musculoskeletal-related inpatient admissions.  |
| Henschke, 2008 <sup>11</sup>   | 973            | Depressive symptoms:<br>Scale for feelings of depression<br>(0-10, higher=worse)                                  | Time to recovery (based on<br>return to previous work<br>status, disability scale 1-5,<br>pain intensity scale 1-6) | Unadjusted HR 0.91 (95% CI 0.89, 0.93)<br>Adjusted HR 0.94 (95% CI 0.91, 0.97)<br>Depressive symptoms were associated with slower recovery   |
| Gurcay, 2009 <sup>12</sup>     | 91             | Depressive symptoms:<br>Beck Depression Inventory (0-<br>63, higher=worse)  | Recovery based on pain<br>(Visual Analogue Scale)<br>and disability (Roland<br>Morris Disability<br>Questionnaire)  | Comparing baseline depressive symptoms in recovered versus non-<br>recovered group (mean, SD) at 2 weeks (p=0.569):<br>Recovered: 13.4 (SD 10.2)<br>Non-recovered: 13.8 (SD 8.5)<br>No significant differences in depressive symptoms when comparing<br>recovered versus non-recovered individuals   |
| Reme, 2009 <sup>13</sup>       | 496            | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression (0-60;<br>higher=worse)                  | Work status (full-duty<br>return to work)   | <u>At 12 weeks:</u><br>Unadjusted OR=0.96 (95% CI 0.94–0.98)<br>Depressive symptoms were associated with lower odds of full-duty<br>return to work.  |
| Smeets, 2009 <sup>14</sup>     | 259            | Depressive symptoms:<br>Depression Anxiety Stress<br>Scales-21 (0-43, higher=worse;<br>for 1 SD of change (SD=8)) | Pain (Numeric Rating<br>Scale), function (Patient-<br>specific Functional Scale)                                    | At 1 year (change for a 1 SD increase of DASS-21):Pain: Unadjusted $\beta$ 0.36 (95% CI 0.17, 0.55), p<0.001   |
| Shaw, 2010 <sup>15</sup>       | 140            | Major depression (Diagnostic<br>Interview Schedule Version III-<br>R)   | Pain (Descriptor<br>Differential Scale)   | At 6 months:<br>Major depression (lifetime): Unadjusted OR 4.99 (95%CI 1.49, 16.76)<br><u>Compared to no lifetime diagnosis:</u><br>Major depression in past 6 months: Unadjusted OR 2.42 (95% CI 0.51,<br>11.39)<br>Major depression in >6 months: Unadjusted OR 12.71 (95% CI 1.51,<br>107.21)<br>Major depression was associated with pain. |

| Study                         | Sample<br>Size | Prognostic Factor Measure   | Outcome Measure   | Key Results   |
|-------------------------------|----------------|---|---|---|
| Hiebert, 2012 <sup>16</sup>   | 253            | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression (0-60;<br>higher=worse)                                | Work status (full duty<br>versus not full duty)   | <u>At 4 weeks:</u><br>Unadjusted OR=1.02 (95% CI 0.99, 1.06)<br><u>At 12 weeks:</u><br>Unadjusted OR=1.01 (95% CI 0.95, 1.08)<br>Depressive symptoms were not associated with work status.  |
| Melloh, 2013 <sup>17-21</sup> | 286            | Depressive symptoms:<br>Zung Self-Rating Depression<br>Scale (20-80, higher=worse)  | Persistent pain (no decrease<br>on Visual Analogue Scale)<br>Pain (SF-MPQ pain<br>subscale)<br>Sickness absence at work or<br>school<br>Disability (Oswestry<br>Disability Index) | Persistent pain at 6 months<br>Adjusted OR 1.03 (95% CI 0.98, 1.08)<br>Pain (SF-MPQ pain subscale)<br>At 3 weeks: Unadjusted $\beta = 0.23$ , p<0.01<br>At 6 weeks: Unadjusted $\beta = 0.17$ , p<0.01<br>Varied results for the association between depressive symptoms and<br>pain.<br>Sickness absence at 3 weeks:<br>Unadjusted OR 1.07 (95% CI 0.98-1.08), SE=0.03<br>Sickness absence at 6 weeks:<br>Unadjusted OR 1.09 (95% CI 1.04, 1.14)<br>Adjusted OR 1.09 (95% CI 1.01, 1.17), SE=0.04<br>Sickness absence at 12 weeks:<br>Unadjusted OR 1.10 (95% CI 1.03, 1.17)<br>Sickness absence at 6 months:<br>Unadjusted OR 1.10 (95% CI 1.04, 1.17)<br>Depressive symptoms were associated with sickness absence.<br>Disability:<br>At 3 weeks: r=0.23, p<0.01<br>At 6 weeks: r=0.43, p<0.01<br>At 6 months: r=0.41, p<0.01<br>Depressive symptoms correlated with disability. |
| Enthoven, 2016 <sup>22</sup>  | 675            | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression (0-60;<br>higher=worse)                                | Pain trajectory (Numeric<br>Rating Scale)   | At 3 years:<br>Intermediate pain trajectory: Unadjusted OR 1.00 (95% CI 0.95, 1.06)<br>p=0.97<br>High pain trajectory: Unadjusted OR 1.01 (95% CI 0.95–1.07) p=0.83<br>Depressive symptoms were not associated with pain trajectory.  |
| Steenstra, 2016 <sup>23</sup> | 113            | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression (0-60,<br>higher=worse; ≥16 as depressive<br>symptoms) | Time on benefits during<br>workers' compensation<br>claim   | Unadjusted HR 0.793 (95% CI 0.54, 1.16), p=0.236<br>Depressive symptoms were not associated with time on benefits during<br>workers' compensation claim.  |

| Study                                | Sample<br>Size | Prognostic Factor Measure   | Outcome Measure   | Key Results   |
|--------------------------------------|----------------|---|---|---|
| Traeger, 2016 <sup>24</sup>          | 2891           | Depressive symptoms:<br>Low Back Pain Screening<br>Questionnaire – depression scale<br>(11-point scale)               | LBP-specific health care<br>utilization (number of<br>primary health care visits)                   | Short-term health care use over 3 months<br>Unadjusted IRR 1.04 (95% CI 1.01, 1.07)<br>Adjusted IRR 1.03 (95% CI 0.99, 1.06)<br>Long-term health care use over 12 months<br>Unadjusted IRR 1.07 (95% CI 1.04, 1.09)<br>Adjusted IRR 1.04 (95% CI 1.02, 1.07)<br>Depressive symptoms were associated with long-term but not short term<br>health care use.   |
| Friedman, 2017 <sup>25</sup>         | 323            | Depressive symptoms:<br>Patient Health Questionnaire-2<br>(dichotomized to "often" or<br>"always" to either question) | Disability (Roland Morris<br>Disability Questionnaire)<br>Pain (ordinal scale "severe<br>to "none") | At 3 months:<br><u>Any functional impairment:</u><br>Unadjusted OR 0.6 (95% CI 0.2-2.3)<br>Adjusted OR 0.9 (95% CI 0.2-3.6)<br><u>Pain:</u><br>Unadjusted OR 0.8 (95% CI 0.2-2.6)<br>Adjusted OR 1.0 (95% CI 0.3-3.7)<br>Depressive symptoms were not associated with disability or pain.   |
| Klyne, 2020 <sup>26</sup>            | 28             | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression (0-60;<br>higher=worse)                      | Pain (Numeric Rating<br>Scale; $\geq$ 7/10 or increase or<br>no change considered non-<br>recovery) | Unrecovered versus recovered LBP at 6 months:<br>Unadjusted OR 1.11 (95% CI 1.00, 1.24), p=0.046<br>Adjusted OR 1.14 (95% CI 0.98, 1.33), p=0.091<br>Individuals with depressive symptoms were more likely to be non-<br>recovered (but not statistically significant).   |
| Acute LBP with radiculopathy         |                |   |   |   |
| Bahar-Ozmedir,<br>2020 <sup>27</sup> | 161            | Depressive symptoms:<br>Hospital Anxiety and Depression<br>Scale-Depression (0-21; ≥11 as<br>depressive symptoms)     | Pain (Numeric Rating<br>Scale), disability (Oswestry<br>Disability Index)                           | $\label{eq:correlation with percent change in pain:} \\ At 1 hour: r=-0.066 p=0.403 \\ At 3-weeks: r=-0.182 p=0.022 \\ At 3-months: r=-0.204 p=0.037 \\ Depressive symptoms correlated with less reduction in pain at 3 weeks \\ and 3 months but not 1 hour post-treatment. No significant differences in \\ depressive symptoms between patients who had successful and failed \\ outcomes (i.e., \geq 50% decrease in pain) (p>0.05).Correlation with percent change in disability:At 3-weeks: r=0.054 p=0.527 \\ At 3-months: r=0.022 p=0.828 \\ Depressive symptoms did not correlate with functional limitations post-treatment.$ |
| Chronic LBP (≥3<br>months' duration) |                |   |   |   |
| Rainville, 1993 <sup>28</sup>        | 72             | Depressive symptoms:<br>Beck Depression Inventory (0-<br>63, higher=worse)  | Pain and disability (Million<br>Visual Analogue Scale)  | Post-treatment           Adjusted r <sup>2</sup> change=0.45, F change=47.13 (p<0.001)  |

| Study                                | Sample<br>Size | Prognostic Factor Measure  | Outcome Measure   | Key Results   |
|--------------------------------------|----------------|--|---|---|
| Harkapaa, 1996 <sup>29</sup>         | 175            | Depressive symptoms:<br>Index of Depression (5 items<br>from Beck Depression Inventory<br>and Cornell Medical Index) | Work status (at work, on<br>sick-leave or pension, other<br>status)   | At 12 months:         F=1.17, not significant         Mean scores in depressive symptoms (not significant):         At work: 7.8         Sick leave pension: 8.1         Other status: 8.4         Depressive symptoms were not associated with work status.  |
| Dionne, 1997 <sup>30</sup>           | 1213           | Depressive symptoms:<br>Symptom Checklist-90-R   | Disability (Roland Morris<br>Disability Questionnaire)  | <u>At 2 years:</u><br>Unadjusted R <sup>2</sup> =0.17, p=0.0001<br>Adjusted R <sup>2</sup> =0.218, p=0.005<br>Depressive symptoms were associated with disability.  |
| Vendrig, 1999 <sup>31</sup>          | 143            | Depressive symptoms:<br>Minnesota Multiphasic<br>Personality Inventory-Depression<br>scale                           | Work status (return to work)  | Mean scores in depressive symptoms comparing return to work status at<br><u>6 months (p=0.419):</u><br>Complete return to work: mean 57.1 (SD 11.3)<br>Incomplete return to work: mean 59.5 (SD 11.7)<br>Depressive symptoms were not associated with return to work.   |
| Michaelson, 2004 <sup>32</sup>       | 167            | Depressive symptoms:<br>Beck Depression Inventory (0-<br>63, higher=worse)   | Pain (Visual Analogue<br>Scale)   | Depressive symptoms demonstrated no predictive value for pain at follow-up (model estimates not reported).  |
| Weidenhammer,<br>2007 <sup>33</sup>  | 4032           | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression Scale<br>(German version)                   | Pain (10-point rating scale),<br>health-related quality of life<br>(SF-36 Physical<br>Component Score, Mental<br>Component Score)                                   | Pain before treatment (P<0.001):         With depressive symptoms: mean 6.3 (SD 1.9)         Without depressive symptoms: mean 5.2 (SD 1.8)         Pain post-treatment:         With depressive symptoms: mean 4.1         Without depressive symptoms: mean 3.2         Compared to patients with depressive symptoms, patients without depressive symptoms had a significantly higher improvement in health-related QOL physical health component, but not mental health component scores or pain improvement. |
| Van Der Hulst,<br>2008 <sup>34</sup> | 163            | Depressive symptoms:<br>Symptom Checklist-90 (16-80,<br>higher=worse)  | Disability (Roland Morris<br>Disability Questionnaire),<br>health-related quality of life<br>(Short Form-36 Physical<br>Component Scale, Mental<br>Component Scale) | After dischargeRMDQ: Adjusted $\beta$ 0.04 (SE 0.05), p>0.05SF-36 PCS: Adjusted $\beta$ -0.09 (SE 0.08), p>0.05SF-36 MCS: Adjusted $\beta$ -0.29 (SE 0.13), p<0.05  |
| Streitberger, 2011 <sup>35</sup>     | 41             | Depressive symptoms:   | Pain (Visual Analogue<br>Scale)   | At 1 year:<br>Adjusted HR 2.97 (95% 1.32, 6.65)   |

| Study                         | Sample<br>Size | Prognostic Factor Measure   | Outcome Measure  | Key Results  |
|-------------------------------|----------------|---|--|--|
|                               |                | Beck Depression Inventory (0-<br>63, higher=worse; >16 as<br>depressive symptoms)                                   |  | Depressive symptoms were associated with shorter duration of pain reduction.   |
| Hicks, 2012 <sup>36</sup>     | 392            | Depressive symptoms:<br>Geriatric Depression Scale (0-15;<br>higher=worse; >5 as depressive<br>symptoms)            | Pain (Numeric Rating<br>Scale)   | <u>At 12 months:</u><br>Adjusted OR 0.47 (95% CI 0.25, 0.89)<br>Depressive symptoms were associated with lower odds of pain<br>reduction.  |
| van Hooff, 2014 <sup>37</sup> | 524            | Depressive symptoms:<br>Zung Self-Rating Depression<br>Scale (20-80, higher=worse)                                  | Disability (Oswestry<br>Disability Index)  | At 1 year:<br><u>Mean (SD) for depressive symptoms</u><br>Disability - success: 56.4 (SD 22.2)<br>Disability- failure 63.7 (SD 19.8)<br>Depressive symptoms were associated with disability.   |
| Scherrer, 2015 <sup>38</sup>  | 355            | Depressive symptoms:<br>Patient Health Questionnaire-2<br>(0-6, higher=worse; ≥3 as<br>depressive symptoms)         | Health care utilization<br>(morphine equivalent<br>opioid dose; high dose>50<br>mg/day)  | Up to 24 months:<br>Unadjusted OR 2.13 (95% CI 1.36, 3.36)<br>Adjusted OR 1.65 (95% CI 0.97, 2.81)<br>Depressive symptoms were associated with higher odds of opioid use,<br>but this was not statistically significant.   |
| Viniol, 2015 <sup>39</sup>    | 484            | Depressive symptoms:<br>Hospital Anxiety and Depression<br>Scale-Depression (0-21,<br>higher=worse)                 | Chronic widespread pain<br>(body pain drawing)   | 1 year:         Adjusted OR 1.01, p=0.87         Depressive symptoms were not associated with the incidence of chronic widespread pain.  |
| Cougot, 2015 <sup>40</sup>    | 217            | Depressive symptoms:<br>Hospital Anxiety and Depression<br>Scale-Depression (0-21,<br>higher=worse)                 | Work status (keeping job<br>after 2 years)   | <u>At 2 years:</u><br>Unadjusted OR 0.92 (95% CI 0.81, 1.05)<br>Depressive symptoms were not associated with work status.  |
| Yarlas, 2016 <sup>41</sup>    | 541            | Depressive symptoms:<br>Short Form-36 Mental Health<br>Domain (0-100, higher=better;<br>≤52 as depressive symptoms) | Pain (Numeric Rating<br>Scale), pain interference<br>(Brief Pain Inventory),<br>health-related quality of life<br>(SF-36), disability<br>(Oswestry Disability Index) | No statistically significant interactions between treatment arm and depression status were observed. The main effect of depression status was not statistically significant on any outcome based on adjusted results except for the BPI Interference subscale ( $P < 0.01$ ), which showed a deficit for depressed patients. |
| Jegan, 2017 <sup>42</sup>     | 484            | Depressive symptoms:<br>Hospital Anxiety and Depression<br>Scale-Depression (0-21,<br>higher=worse)                 | Disability<br>(Multidimensional Pain<br>Inventory-Disability)  | At 1 year:<br>Adjusted $\beta$ 0.83 (95% CI 0.01, 1.64)<br>Depressive symptoms were associated with disability.  |
| Nordeman, 2017 <sup>43</sup>  | 130            | Depressive symptoms:<br>Hospital Anxiety and Depression<br>Scale-Depression (0-21,<br>higher=worse)                 | Disability (Roland Morris<br>Disability Questionnaire)   | <u>At 2 years:</u><br>Activity limitation on RMDQ: r=0.37 p=0.000025<br>Percent change in RMDQ: r=-0.11 p=0.24<br>Depressive symptoms were correlated with activity limitation but not<br>percent change on RMDQ.  |
| Nordstoga, 2017 <sup>44</sup> | 7523           | Depressive symptoms:  | Recovery (no pain or stiffness in muscles or   | Women:<br>Unadjusted RR 0.83   |

| Study                          | Sample<br>Size | Prognostic Factor Measure   | Outcome Measure   | Key Results  |
|--------------------------------|----------------|---|---|--|
|                                |                | Hospital Anxiety and Depression<br>Scale-Depression (0-21,<br>higher=worse; ≥8 as depressive<br>symptoms) | joints for ≥3 consecutive<br>months)  | Adjusted RR 0.90 (95% CI 0.76, 1.07)<br><u>Men:</u><br>Unadjusted RR 0.85<br>Adjusted RR 0.85 (95% CI 0.73, 0.99)<br>Depressive symptoms were associated with reduced risk of recovery in<br>men and women (not statistically significant in women).   |
| Dengler, 2018 <sup>45</sup>    | 101            | Depressive symptoms:<br>Zung Self-Rating Depression<br>Scale (20-80, higher=worse)                        | Health care utilization<br>(opioid use)   | Conservative management group:<br>At 6 months: Adjusted OR 1.00 (95% CI 0.87, 1.20)<br>Depressive symptoms were not associated with opioid use.  |
| Glattacker, 2018 <sup>46</sup> | 214            | Depressive symptoms:<br>Hospital Anxiety and Depression<br>Scale-Depression (0-21,<br>higher=worse)       | Health-related quality of<br>life (Short Form-12<br>Physical Component Score,<br>Mental Component Score)                  | SF-12 Physical Component Score:Depressive symptoms did not reach threshold of significance (based on<br>unadjusted results).SF-12 Mental Component Score:End of rehab: Adjusted $\beta$ =-0.703 p=0.0026-months post-rehab: Adjusted $\beta$ =-0.989 p=0.007Depressive symptoms were associated with health-related quality of life<br>mental component but not physical component.  |
| Halonen, 2019 <sup>47</sup>    | 5740           | Depressive symptoms:<br>SCL-Core Depression scale (≥17<br>as major depressive symptoms)                   | Persistent pain and<br>functional limitations<br>(single question)  | <u>Up to 6 years of follow-up:</u><br>Adjusted RR 1.09 (95% CI 1.05, 1.14)<br>Depressive symptoms were associated with persistent pain and<br>functional limitations.  |
| Demarchi, 2019 <sup>48</sup>   | 92             | Depressive symptoms:<br>Beck Depression Inventory (0-<br>63, higher=worse)                                | Pain (Numeric Rating<br>Scale), disability (Roland<br>Morris Disability<br>Questionnaire)                                 | At 6 months:Pain: Adjusted $\beta$ 0.09 (95% CI 0.02, 0.16) p=0.01At 6 months:Functional limitations: Adjusted $\beta$ 0.13 (95% CI -0.03, 0.29) p=0.10Depressive symptoms were associated with pain but not disability.   |
| Oliveira, 2019 <sup>49</sup>   | 284            | Depressive symptoms:<br>Hospital Anxiety and Depression<br>Scale-Depression (0-21,<br>higher=worse)       | Pain (Brief Pain Inventory);<br>pain, disability, and<br>satisfaction (Shortened<br>Treatment Outcomes in<br>Pain Survey) | At 1 year:BPI Severity: Adjusted β 0.05 (95% CI –0.007, 0.11)BPI Interference: Adjusted β 0.31 (95% CI 0.20, 0.43)S-TOPS:Pain symptoms: Adjusted β 0.58 (95% CI –0.04, 1.19)Physical disability–lower body: Adjusted β 2.41 (95% CI 0.65, 4.16)Physical disability–upper body: Adjusted β 0.36 (95% CI –0.08, 0.79)Family/social disability: Adjusted β 1.94 (95% CI 1.27, 2.60)Role emotional disability: Adjusted β 1.75 (95% CI 0.88, 2.63)Patient satisfaction with care: Adjusted β -0.23 (95% CI –0.79, 0.34)Patient satisfaction with outcomes: Adjusted β -0.46 (95% CI –1.07, 0.15)Depressive symptoms were associated with disability, but notsatisfaction, and varied with pain measures. |
| Page, 2019 <sup>50</sup>       | 686            | Depressive symptoms:  | Pain severity (based on<br>Numeric Rating Scale and 2   | <u>At 1 year:</u>  |

| Study                          | Sample<br>Size | Prognostic Factor Measure  | Outcome Measure  | Key Results   |
|--------------------------------|----------------|--|--|---|
|                                |                | Beck Depression Inventory (0-<br>63, higher=worse)   | items from Brief Pain<br>Inventory-10: enjoyment of<br>life and general activities)  | Adjusted $\beta$ (intercept) 0.044, SE 0.011, p<0.001; $\beta$ (slope) –0.002, SE 0.008, p=0.773<br>Depressive symptoms were not associated with rates of change in pain severity.  |
| Imagama, 2020 <sup>51</sup>    | 474            | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression (0-60,<br>higher=worse)   | Health-related quality of<br>life (Short Form-8 Mental<br>Component Score)   | At 6 months:<br>Adjusted β 2.64 (95% CI 1.53, 3.75) p<0.001<br>Depressive symptoms were associated with improved health-related<br>quality of life.   |
| Ranger, 2020 <sup>52</sup>     | 633            | Depressive symptoms:<br>Screening Index for Spine Data<br>Registry-depression scale (0-10,<br>higher=worse; ≥7 as depressive<br>symptoms)      | Pain (Numeric Rating<br>Scale), disability (Roland<br>Morris Disability<br>Questionnaire)  | At 1 year: <u>Pain:</u> Unadjusted RR 2.07 (95% CI 1.68, 2.57) <.001  |
| Zackova, 2020 <sup>53</sup>    | 413            | Depressive symptoms:<br>Beck Depression Inventory [0-<br>63, higher=worse; 0–9 (normal);<br>10–18 (mild); 19–29 (moderate);<br>30–63 (severe)] | Pain (Numeric Rating<br>Scale), health-related<br>quality of life (Short Form-<br>36 Physical Component<br>Scale, Mental Component<br>Scale) | At 6 months:<br><u>PCS improvement (yes vs no)</u><br>BDI Moderate versus mild: Adjusted OR 0.85 (95% CI 0.32, 2.22),<br>p=0.74<br>BDI Severe versus mild: Adjusted OR 1.37 (95% CI 0.35, 5.31), p=0.65<br><u>MCS improvement (yes vs no)</u><br>BDI Moderate versus mild: Adjusted OR 0.14 (95% CI 0.03, 0.57),<br>p=0.01<br>BDI Severe versus mild: Adjusted OR 0.04 (95% CI 0.01, 0.27),<br>p=0.001<br><u>Improvement based on decrease in NRS (p=0.13)</u><br>BDI Mild n (%): improved 5 (55.6) vs not improved 61 (35.5)<br>BDI Moderate n (%): improved 2 (22.2) vs not improved 69 (40.1),<br>BDI Severe n (%): improved 2 (22.2) vs not improved 42 (24.4)<br>Depressive symptoms were associated with lower odds of improvement<br>in health-related quality of life mental component but not physical<br>component or pain reduction. |
| Chronic LBP with radiculopathy |                |  |  |   |
| Lubelski, 2015 <sup>54</sup>   | 1346           | Depressive symptoms:<br>Patient Health Questionnaire-9<br>(0-27, higher=worse)   | Quality of life (EuroQol-<br>5D), health care utilization<br>(surgery)   | At 4 months:<br>Quality of life (any improvement): Adjusted OR 0.95 (95% CI 0.90,<br>1.00)<br>Quality of life (≥0.1 point improvement): Adjusted OR 0.92 (95% CI<br>0.87, 0.98)<br>Surgery: Adjusted OR 1.03 (95% CI 0.98, 1.08)<br>Time to surgery: Adjusted HR 1.02 (95% CI 0.98, 1.06)   |

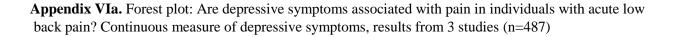
| Study                       | Sample<br>Size | Prognostic Factor Measure   | Outcome Measure  | Key Results   |
|-----------------------------|----------------|---|--|---|
|                             |                |   |  | Depressive symptoms were associated with reduced quality of life but not surgery.   |
| Mixed duration              |                |   |  |   |
| Newell, 2007 <sup>55</sup>  | 788            | Depressive symptoms:<br>Bournemouth Questionnaire   | Recovery (Global<br>Impression of Change)  | Acute LBP <4 weeks' duration n=241:At 4 weeks: Unadjusted OR 0.31 (95% CI 0.14, 0.69)At 12 weeks: not significantAcute LBP <4 weeks' duration, recurring  |
| Grotle, 2010 <sup>56</sup>  | 926            | Depressive symptoms:<br>Hospital Anxiety and Depression<br>Scale-Depression (0-21,<br>higher=worse)   | Disability (Roland Morris<br>Disability Questionnaire)   | 1 year:Acute/subacute LBP (<3 months' duration) n=258:  |
| Adnan, 2017 <sup>57</sup>   | 565            | Depressive symptoms:<br>Beck Depression Inventory (0-<br>63, higher=worse)                            | Favourable outcome: 30%<br>reduction in both disability<br>(Oswestry Disability Index)<br>and pain (Numeric Rating<br>Scale) | Post-treatment:         Acute LBP (<14 weeks' duration) n=153:  |
| Other <sup>a</sup>          |                |   |  |   |
| Cherkin, 1996 <sup>58</sup> | 219            | Depressive symptoms:<br>Symptom Checklist-90<br>(reference: <moderately<br>depressed)</moderately<br> | Tolerability of symptom<br>severity (poor outcome:<br>mostly dissatisfied,<br>unhappy, or terrible)                          | In participants with index visit for low back pain:<br>7 weeks: Adjusted OR 2.1 (95% CI 1.4, 3.4)<br>1 year: Adjusted OR 2.3 (95% CI 1.4, 3.6)<br>Depressive symptoms were associated with poor symptom satisfaction. |
| Fransen, 2002 <sup>59</sup> | 854            | Depressive symptoms:<br>General Health<br>Questionnaire (severe<br>depression)                        | Compensation claim status<br>(payments) for low back<br>pain   | New occupational claim for LBP:<br>At 12 weeks:<br>Adjusted OR=2.47 (95% CI 1.66, 3.67)   |

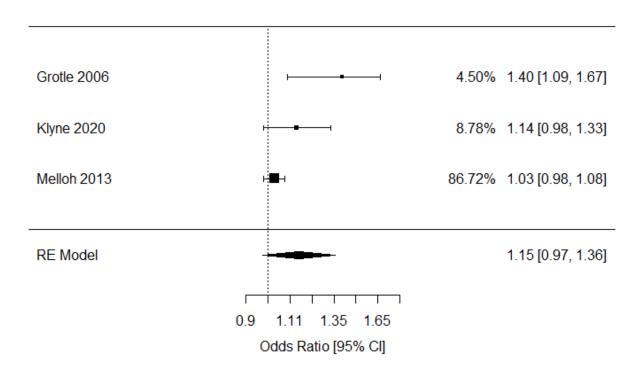
| Study                        | Sample<br>Size | Prognostic Factor Measure  | Outcome Measure  | Key Results  |
|------------------------------|----------------|--|--|--|
|                              |                |  |  | Depressive symptoms were associated with compensation claims for low back pain.  |
| Tubach, 2004 <sup>60</sup>   | 622            | Depressive symptoms:<br>Langner's Screening<br>Questionnaire (depression score;<br>score dichotomized to high<br>versus low) | Persistence of sciatica<br>(single question)   | LBP with radiculopathy ≥1 day during previous year:<br>At 2 years:<br>Adjusted OR 0.87 (95% CI 0.55, 1.37)<br>Depressive symptoms were not associated with persistence of sciatica.  |
| Campello, 2006 <sup>61</sup> | 67             | Depressive symptoms:<br>Symptom Checklist-90-R   | Work status: time to work retention  | Off/restricted duty >8 weeks:<br>At 2 years:<br>Unadjusted HR 0.72 (95% CI 0.22, 2.44)<br>Multivariate analysis: not statistically associated with work retention<br>Depressive symptoms were not associated with work status.   |
| Scheele, 2013 <sup>62</sup>  | 675            | Depressive symptoms:<br>Center for Epidemiological<br>Studies Depression (0-60;<br>higher=worse)                             | Non-recovery (Global<br>Perceived Effect)  | In participants with index visit for low back pain:<br>At 3 months:<br>Unadjusted OR 1.1, p<0.01<br>Depressive symptoms were associated with non-recovery.   |
| Fischer, 2014 <sup>63</sup>  | 395            | Depressive symptoms:<br>Zung Self-Rating Depression<br>Scale (20-80, higher=worse)   | Quality of life (Freiburger<br>Persönlichkeitsinventar<br>subjective well-being), pain<br>(Visual Analogue Scale),<br>work status (return to work) | LBP with sick leave:<br><u>At 6 months:</u><br>Lower quality of life: Adjusted OR 1.48 p<0.0001<br>Pain reduction: Adjusted OR 0.93 (not significant)<br>Return to work: Adjusted OR 0.86 (not significant)<br>Depressive symptoms were associated with quality of life, but not<br>return-to-work or pain outcomes.   |
| Kerr, 2015 <sup>64</sup>     | 392            | Depression (history of depression)   | Disability (Roland Morris<br>Disability Questionnaire)   | Among participants with lumbar radiculopathy ≥6 weeks' duration:<br>Mean change scores from baseline (non-operative treatment group) at 8<br>years follow-up:<br>Without depressive symptoms: -25.4 (SE 0.7)<br>With depressive symptoms: -21.6 (SE 1.8)<br>Depressive symptoms were associated with disability.   |
| Kim, 2017 <sup>65</sup>      | 161            | Depressive symptoms:<br>Zung Self-Rating Depression<br>Scale (20-80, higher=worse)   | Disability (Oswestry<br>Disability Index)  | LBP with radiculopathy with non-response to care:<br><u>At 12 months:</u><br>Adjusted Beta 0.465 (95% CI 0.146, 0.784) p=0.005<br>Depressive symptoms were associated with disability.   |
| Hartvigsen, 2020 –           | 947            | Depressive symptoms:<br>Major Depression Inventory (0-<br>50, higher=worse; > 19 as<br>depressive symptoms)                  | Pain (Numeric Rating<br>Scale), disability (Roland<br>Morris Disability<br>Questionnaire)  | Index health care visit for LBP (chiropractic patients):           Pain:           At 2 weeks: Unadjusted OR 2.71 (95% CI 1.65, 4.43)           At 3 months: Unadjusted OR 3.06 (95% CI 1.91, 4.91)           At 1 year: Unadjusted OR 2.32 (95% CI 1.41, 3.81)           Disability:           At 2 weeks: Unadjusted OR 3.44 (95% CI 2.07, 5.70)           At 3 months: Unadjusted OR 2.57 (95% CI 1.64, 4.05)           At 1 year: Unadjusted OR 1.59 (95% CI 0.96, 2.64) |

| Study | Sample<br>Size | Prognostic Factor Measure | Outcome Measure | Key Results   |
|-------|----------------|---------------------------|-----------------|---|
|       |                |                           |                 | Pain as a categorical variable at 3 months:                       |
|       |                |                           |                 | Medium: Unadjusted RR 1.96 (95% CI 1.04-3.71)                     |
|       |                |                           |                 | High: Unadjusted RR 5.22 (95% CI 2.62-10.4)                       |
|       |                |                           |                 | Disability as a categorical variable at 3 months (reference=low): |
|       |                |                           |                 | Medium: Unadjusted RR 1.92 (95% CI 1.03-3.59)                     |
|       |                |                           |                 | High: Unadjusted RR 3.50 (95% CI 1.92-6.38)                       |
|       |                |                           |                 | Depressive symptoms were associated with pain and disability.     |

CI – confidence interval; HR – hazard ratio; LBP – low back pain; OR – odds ratio; RR – relative risk; RMDQ – Roland Morris Disability Questionnaire; SD – standard deviation; SE – standard error; VAS – visual analogue scale

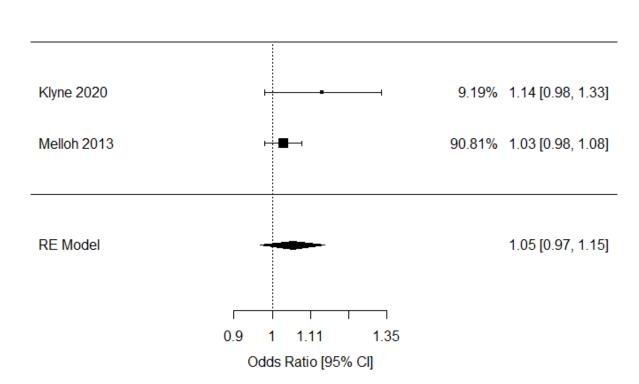
<sup>a</sup>Other LBP populations as described under the key results column





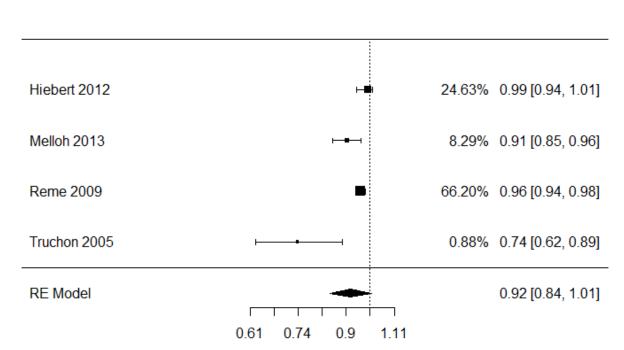
RE – random effects meta-analysis; tau<sup>2</sup>=0.02, I<sup>2</sup>=78.67%\*

\* I<sup>2</sup> <75% when excluding one high risk of bias study (Grotle 2006; see Appendix VIb)



**Appendix VIb.** Forest plot: Are depressive symptoms associated with pain in individuals with acute low back pain? Continuous measure of depressive symptoms, results from 2 studies (n=314)

RE - random effects meta-analysis; tau<sup>2</sup><0.01, I<sup>2</sup>=35.09%

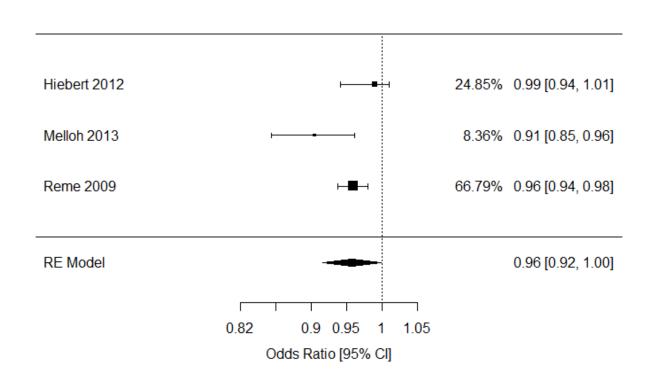


**Appendix VIIa.** Forest plot: Are depressive symptoms associated with work participation in individuals with acute low back pain? Continuous measure of depressive symptoms, results from 4 studies (n=1,356)

Odds Ratio [95% CI]

RE - random effects meta-analysis; tau<sup>2</sup>=0.01, I<sup>2</sup>=93.46%

\* I<sup>2</sup> decreased when excluding one high risk of bias studies (Truchon 2005; see Appendix VIIb)

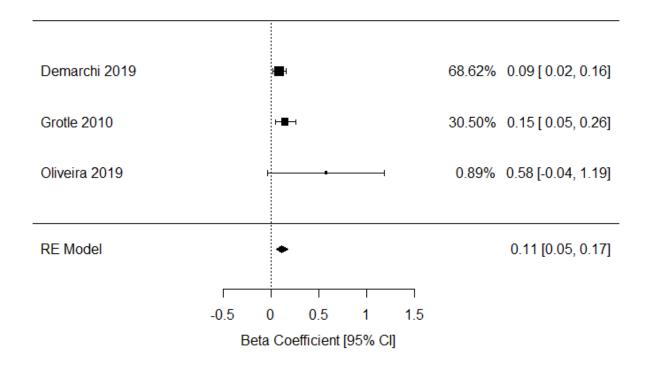


**Appendix VIIb.** Forest plot: Are depressive symptoms associated with work participation in individuals with acute low back pain? Continuous measure for depressive symptoms, results from 3 studies (n=1,035)

RE – random effects meta-analysis; tau<sup>2</sup><0.01, I<sup>2</sup>=75.70%\*

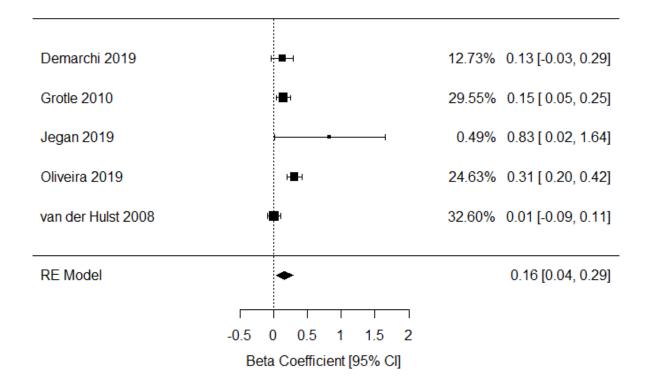
\*To explore potential sources of heterogeneity, we further excluded one moderate risk of bias study from the meta-analysis, which produced similar results (2 studies, 749 participants, OR=0.97, 95% CI 0.94-1.00,  $I^2$ =56.27%)

**Appendix VIII.** Forest plot: Are depressive symptoms associated with pain in individuals with chronic low back pain? Continuous measure of depressive symptoms, results from 3 studies (n=2,902)



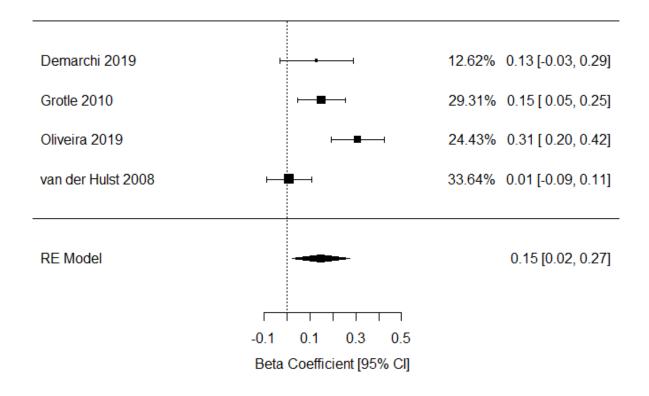
RE – random effects meta-analysis; tau<sup>2</sup>=0, I<sup>2</sup>=1.04%

**Appendix IXa.** Forest plot: Are depressive symptoms associated with disability in individuals with chronic low back pain? Continuous measure of depressive symptoms, results from 5 studies (n=3,549)



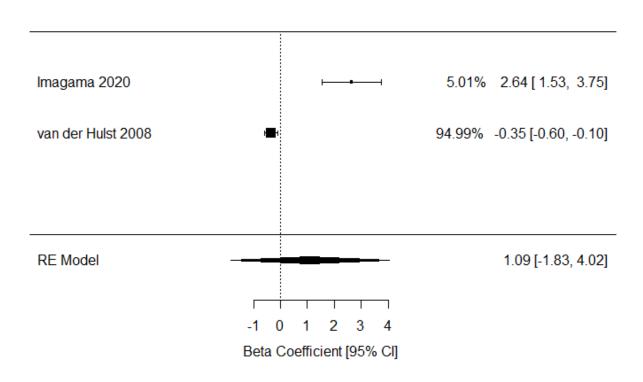
RE – random effects meta-analysis; tau<sup>2</sup>=0.01, I<sup>2</sup>=74.69%

**Appendix IXb.** Forest plot: Are depressive symptoms associated with disability in individuals with chronic low back pain? Continuous measure of depressive symptoms, results from 4 studies (n=3,065)



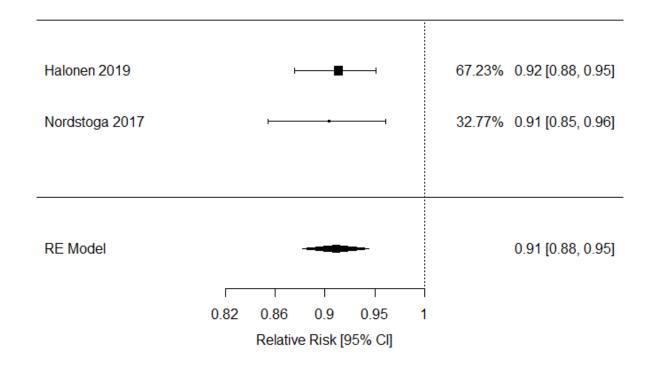
RE – random effects meta-analysis; tau<sup>2</sup>=0.01, I<sup>2</sup>= 78.70%

**Appendix X.** Forest plot: Are depressive symptoms associated with the mental component of healthrelated quality of life in individuals with chronic low back pain? Continuous measure of depressive symptoms, results from 2 studies (n=637)



RE - random effects meta-analysis; tau<sup>2</sup>=4.30, I<sup>2</sup>=96.22%\*

\*Could not adequately explore sources of heterogeneity due to a limited number of studies



**Appendix XI.** Forest plot: Are depressive symptoms associated with recovery in individuals with chronic low back pain? Dichotomous measure of depressive symptoms, results from 2 studies (n=13,263)

RE - random effects meta-analysis; tau<sup>2</sup>=0, I<sup>2</sup>=0%

Appendix XIIa. Summary of the domains of the Quality in Prognosis Studies (QUIPS) tool\*

| Biases  | Issues to consider for judging overall rating of "Risk of bias"   |  |  |  |  |
|---|---|--|--|--|--|
| 1. Study Participation  | Goal: To judge the risk of selection bias (likelihood that relationship between <i>PF</i> and <i>outcome</i> is different for participants and eligible non-participants).                            |  |  |  |  |
| Source of target population                                     | The source population or population of interest is adequately described for key characteristics.  |  |  |  |  |
| Method used to identify population                              | The sampling frame and recruitment are adequately described, including methods to identify the samp sufficient to limit potential bias (number and type used, e.g., referral patterns in health care) |  |  |  |  |
| Recruitment period  | Period of recruitment is adequately described   |  |  |  |  |
| Place of recruitment  | Place of recruitment (setting and geographic location) are adequately described   |  |  |  |  |
| Inclusion and exclusion criteria                                | Inclusion and exclusion criteria are adequately described (e.g., including explicit diagnostic criteria or "zero time" description).  |  |  |  |  |
| Adequate study participation                                    | There is adequate participation in the study by eligible individuals  |  |  |  |  |
| Baseline characteristics  | The baseline study sample (i.e., individuals entering the study) is adequately described for key characteristics.   |  |  |  |  |
| Summary Study participation                                     | The study sample represents the population of interest on key characteristics, sufficient to limit potential bias of the observed relationship between PF and outcome.                                |  |  |  |  |
| 2. Study Attrition  | Goal: To judge the risk of attrition bias (likelihood that relationship between <i>PF</i> and <i>outcome</i> are different for completing and non-completing participants).                           |  |  |  |  |
| Proportion of baseline sample available for analysis            | Response rate (i.e., proportion of study sample completing the study and providing outcome data) is adequate.   |  |  |  |  |
| Attempts to collect information on participants who dropped out | Attempts to collect information on participants who dropped out of the study are described.   |  |  |  |  |
| Reasons and potential impact of subjects lost to follow-up      | Reasons for loss to follow-up are provided.   |  |  |  |  |
| Outcome and prognostic factor                                   | Participants lost to follow-up are adequately described for key characteristics.  |  |  |  |  |
| information on those lost to follow-up                          | There are no important differences between key characteristics and outcomes in participants who completed the study and those who did not.  |  |  |  |  |

| Study Attrition Summary                         | Loss to follow-up (from baseline sample to study population analyzed) is not associated with key characteristics (i.e., the study data adequately represent the sample) sufficient to limit potential bias to the observed relationship between PF and outcome.         Goal: To judge the risk of measurement bias related to how PF was measured (differential measurement of PF related to the level of outcome). |  |  |  |  |
|---|--|--|--|--|--|
| 3. Prognostic Factor Measurement                |  |  |  |  |  |
| Definition of the PF                            | A clear definition or description of 'PF' is provided (e.g., including dose, level, duration of exposure, and clear specification of the method of measurement).   |  |  |  |  |
| Valid and Reliable Measurement of<br>PF         | Method of PF measurement is adequately valid and reliable to limit misclassification bias (e.g., may include relevant outside sources of information on measurement properties, also characteristics, such as blind measurement and limited reliance on recall).<br>Continuous variables are reported or appropriate cut-points (i.e., not data-dependent) are used.   |  |  |  |  |
| Method and Setting of PF<br>Measurement         | The method and setting of measurement of PF is the same for all study participants.  |  |  |  |  |
| Proportion of data on PF available for analysis | Adequate proportion of the study sample has complete data for PF variable.   |  |  |  |  |
| Method used for missing data                    | Appropriate methods of imputation are used for missing 'PF' data.  |  |  |  |  |
| PF Measurement Summary                          | <i>PF</i> is adequately measured in study participants to sufficiently limit potential bias.   |  |  |  |  |
| 4. Outcome Measurement                          | Goal: To judge the risk of bias related to the measurement of outcome (differential measurement of outcome related to the baseline level of PF).   |  |  |  |  |
| Definition of the Outcome                       | A clear definition of outcome is provided, including duration of follow-up and level and extent of the outcome construct.  |  |  |  |  |
| Valid and Reliable Measurement of<br>Outcome    | The method of outcome measurement used is adequately valid and reliable to limit misclassification bias (e.g., may include relevant outside sources of information on measurement properties, also characteristics, such as blind measurement and confirmation of outcome with valid and reliable test).   |  |  |  |  |
| Method and Setting of Outcome<br>Measurement    | The method and setting of outcome measurement is the same for all study participants.  |  |  |  |  |
| Outcome Measurement Summary                     | <i>Outcome of interest</i> is adequately measured in study participants to sufficiently limit potential bias.  |  |  |  |  |
| 5. Study Confounding                            | Goal: To judge the risk of bias due to confounding (i.e. the effect of PF is distorted by another factor that is related to PF and outcome).   |  |  |  |  |

| Important Confounders Measured                   | All important confounders, including treatments, are measured.   |  |  |  |  |
|--|--|--|--|--|--|
| Definition of the confounding factor             | Clear definitions of the important confounders measured are provided (e.g., including dose, level, and duration of exposures).             |  |  |  |  |
| Valid and Reliable Measurement of                | Measurement of all important confounders is adequately valid and reliable (e.g., may include relevant                                      |  |  |  |  |
| Confounders                                      | outside sources of information on measurement properties, also characteristics, such as blind measurement and limited reliance on recall). |  |  |  |  |
| Method and Setting of Confounding<br>Measurement | The method and setting of confounding measurement are the same for all study participants.   |  |  |  |  |
| Method used for missing data                     | Appropriate methods are used if imputation is used for missing confounder data.  |  |  |  |  |
| Appropriate Accounting for                       | Important potential confounders are accounted for in the study design (e.g., matching for key variables,                                   |  |  |  |  |
| Confounding                                      | stratification, or initial assembly of comparable groups).   |  |  |  |  |
|  | Important potential confounders are accounted for in the analysis (i.e., appropriate adjustment).  |  |  |  |  |
| Study Confounding Summary                        | Important potential confounders are appropriately accounted for, limiting potential bias with  |  |  |  |  |
|  | respect to the relationship between <i>PF</i> and <i>outcome</i> .   |  |  |  |  |
| 6. Statistical Analysis and<br>Reporting         | Goal: To judge the risk of bias related to the statistical analysis and presentation of results.   |  |  |  |  |
| Presentation of analytical strategy              | There is sufficient presentation of data to assess the adequacy of the analysis.   |  |  |  |  |
| Model development strategy                       | The strategy for model building (i.e., inclusion of variables in the statistical model) is appropriate and is                              |  |  |  |  |
|  | based on a conceptual framework or model.  |  |  |  |  |
|  | The selected statistical model is adequate for the design of the study.  |  |  |  |  |
| Reporting of results                             | There is no selective reporting of results.  |  |  |  |  |
| Statistical Analysis and                         | The statistical analysis is appropriate for the design of the study, limiting potential for  |  |  |  |  |
| Presentation Summary                             | presentation of invalid or spurious results.   |  |  |  |  |

PF - prognostic factor

\*Each domain is rated as high, moderate, or low risk of bias considering the prompting items.

| Study                             | Domain 1 –<br>Study<br>Participation | Domain 2 –<br>Study Attrition | Domain 3 –<br>Prognostic<br>Factor<br>Measurement | Domain 4 –<br>Outcome<br>Measurement | Domain 5 –<br>Study<br>Confounding <sup>b</sup> | Domain 6 –<br>Statistical<br>Analysis and<br>Reporting |
|-----------------------------------|--------------------------------------|-------------------------------|---|--------------------------------------|---|--|
| Lehmann, 1993 <sup>1</sup>        | Moderate                             | Low                           | Low   | Moderate                             | High  | Low  |
| Rainville, 1993 <sup>28</sup>     | Low                                  | Moderate                      | Low   | Moderate                             | High  | Low  |
| Gatchel, 1995 <sup>2</sup>        | High                                 | Low                           | Low   | Moderate                             | High  | Moderate   |
| Cherkin, 1996 <sup>58</sup>       | Moderate                             | Low                           | Low   | Moderate                             | Moderate  | Low  |
| Harkapaa, 1996 <sup>29</sup>      | High                                 | Moderate                      | Moderate  | Low                                  | Low   | Low  |
| Dionne, 1997 <sup>30</sup>        | Moderate                             | Low                           | Low   | Low                                  | High  | Low  |
| Epping-Jordan, 1998 <sup>3</sup>  | Low                                  | Moderate                      | Low   | Low                                  | Moderate  | Low  |
| Vendrig, 1999 <sup>31</sup>       | Low                                  | Low                           | Low   | Moderate                             | High  | Moderate   |
| Fritz, 2001 <sup>4</sup>          | Moderate                             | Low                           | Low   | Moderate                             | High  | Low  |
| Fransen, 2002 <sup>59</sup>       | High                                 | Low                           | Low   | Low                                  | Moderate  | Low  |
| Cassidy, 2003 <sup>5</sup>        | Low                                  | Moderate                      | Low   | Low                                  | Low   | Low  |
| Michaelson, 2004 <sup>32</sup>    | Low                                  | Low                           | Low   | Low                                  | Moderate  | Moderate   |
| Tubach, 2004 <sup>60</sup>        | Moderate                             | Moderate                      | Moderate  | Moderate                             | Moderate  | Moderate   |
| Sieben, 2005 <sup>6</sup>         | Moderate                             | Moderate                      | Low   | Low                                  | High  | Low  |
| Truchon, 2005 <sup>7</sup>        | High                                 | Moderate                      | Low   | Moderate                             | High  | Low  |
| Campello, 2006 <sup>61</sup>      | Moderate                             | Low                           | Low   | Moderate                             | Moderate  | Low  |
| Grotle, 2006 <sup>8</sup>         | High                                 | Moderate                      | Moderate  | Low                                  | Moderate  | Low  |
| Patel, 2006 <sup>9</sup>          | High                                 | Moderate                      | Low   | Moderate                             | High  | Moderate   |
| Ritzwoller, 2006 <sup>10</sup>    | Moderate                             | Moderate                      | Moderate  | Moderate                             | Low   | Low  |
| Newell, 2007 <sup>55</sup>        | Moderate                             | High                          | Low   | Low                                  | High  | Moderate   |
| Weidenhammer, 2007 <sup>33</sup>  | Low                                  | Moderate                      | Low   | Low                                  | High  | Moderate   |
| Henschke, 2008 <sup>11</sup>      | Low                                  | Low                           | Moderate  | Low                                  | Low   | Moderate   |
| Van Der Hulst, 2008 <sup>34</sup> | Moderate                             | Moderate                      | Low   | Low                                  | Moderate  | Low  |
| Gurcay, 2009 <sup>12</sup>        | Moderate                             | Low                           | Low   | Low                                  | High  | Low  |

Appendix XIIb. Risk of bias assessment of studies using the Quality in Prognosis Studies (QUIPS) tool

| Study                            | Domain 1 –<br>Study<br>Participation | Domain 2 –<br>Study Attrition | Domain 3 –<br>Prognostic<br>Factor<br>Measurement | Domain 4 –<br>Outcome<br>Measurement | Domain 5 –<br>Study<br>Confounding <sup>b</sup> | Domain 6 –<br>Statistical<br>Analysis and<br>Reporting |
|----------------------------------|--------------------------------------|-------------------------------|---|--------------------------------------|---|--|
| Reme, 2009 <sup>13</sup>         | Moderate                             | Low                           | Moderate  | Low                                  | Low   | Moderate   |
| Smeets, 2009 <sup>14</sup>       | Moderate                             | Low                           | Low   | Low                                  | High  | Low  |
| Grotle, 2010 <sup>56</sup>       | Moderate                             | Moderate                      | Low   | Low                                  | Moderate  | Low  |
| Shaw, 2010 <sup>15</sup>         | Low                                  | Moderate                      | Low   | Moderate                             | High  | Moderate   |
| Streitberger, 2011 <sup>35</sup> | Low                                  | Moderate                      | Low   | Low                                  | Moderate  | Moderate   |
| Hicks, 2012 <sup>36</sup>        | Moderate                             | Moderate                      | Low   | Low                                  | Moderate  | Moderate   |
| Hiebert, 2012 <sup>16</sup>      | Low                                  | Low                           | Low   | Moderate                             | High  | Low  |
| Melloh, 2013 <sup>17-21</sup>    | Moderate                             | Moderate                      | Low   | Low                                  | Moderate  | Low  |
| Scheele, 2013 <sup>62</sup>      | Moderate                             | Low                           | Low   | Low                                  | High  | Moderate   |
| Fischer, 2014 <sup>63</sup>      | Moderate                             | Moderate                      | Low   | Moderate                             | Moderate  | Moderate   |
| van Hooff, 2014 <sup>37</sup>    | Moderate                             | Low                           | Low   | Low                                  | High  | Moderate   |
| Scherrer, 2015 <sup>38</sup>     | Moderate                             | Moderate                      | Low   | Moderate                             | Low   | Low  |
| Viniol, 2015 <sup>39</sup>       | Moderate                             | Moderate                      | Low   | Low                                  | Low   | Low  |
| Cougot, 2015 <sup>40</sup>       | Moderate                             | Moderate                      | Low   | Moderate                             | High  | Low  |
| Kerr, 2015 <sup>64</sup>         | Moderate                             | Moderate                      | Low   | Low                                  | Low   | Moderate   |
| Lubelski, 2015 <sup>54</sup>     | Moderate                             | High                          | Low   | Moderate                             | Low   | Low  |
| Enthoven, 2016 <sup>22</sup>     | Moderate                             | Low                           | Low   | Low                                  | High  | Moderate   |
| Steenstra, 2016 <sup>23</sup>    | Low                                  | Low                           | Low   | Low                                  | High  | Low  |
| Traeger, 2016 <sup>24</sup>      | Moderate                             | Low                           | Low   | Moderate                             | Low   | Low  |
| Yarlas, 2016 <sup>41</sup>       | High                                 | High                          | Low   | Low                                  | High  | Moderate   |
| Adnan, 2017 <sup>57</sup>        | Moderate                             | Low                           | Low   | Moderate                             | Moderate  | Moderate   |
| Friedman, 2017 <sup>25</sup>     | Moderate                             | Low                           | Low   | Moderate                             | Low   | Low  |
| Jegan, 2017 <sup>42</sup>        | Moderate                             | High                          | Low   | Moderate                             | Low   | Low  |
| Nordeman, 2017 <sup>43</sup>     | Moderate                             | Low                           | Low   | Low                                  | High  | Moderate   |
| Nordstoga, 201744                | Moderate                             | Moderate                      | Low   | Moderate                             | Low   | Low  |
| Dengler, 201845                  | Moderate                             | Low                           | Low   | Moderate                             | Moderate  | Low  |

| Study                             | Domain 1 –<br>Study<br>Participation | Domain 2 –<br>Study Attrition | Domain 3 –<br>Prognostic<br>Factor<br>Measurement | Domain 4 –<br>Outcome<br>Measurement | Domain 5 –<br>Study<br>Confounding <sup>b</sup> | Domain 6 –<br>Statistical<br>Analysis and<br>Reporting |
|-----------------------------------|--------------------------------------|-------------------------------|---|--------------------------------------|---|--|
| Oliveira, 2019 <sup>49</sup>      | Moderate                             | Moderate                      | Low   | Low                                  | Low   | Low  |
| Page, 2019 <sup>50</sup>          | Moderate                             | Low                           | Low   | Low                                  | Low   | Low  |
| Bahar-Ozmedir, 2020 <sup>27</sup> | Moderate                             | High                          | Low   | Moderate                             | High  | Low  |
| Hartvigsen, 2020 <sup>66</sup>    | Low                                  | Moderate                      | Low   | Low                                  | Low   | Low  |
| Imagama, 2020 <sup>51</sup>       | Moderate                             | Moderate                      | Low   | Low                                  | Low   | Low  |
| Klyne, 2020 <sup>26</sup>         | Moderate                             | Low                           | Low   | Low                                  | Moderate  | Low  |
| Ranger, 2020 <sup>52</sup>        | Low                                  | Moderate                      | Low   | Low                                  | Low   | Low  |
| Zackova, 2020 <sup>53</sup>       | Low                                  | Moderate                      | Low   | Moderate                             | Moderate  | Low  |

<sup>a</sup>Domains were assessed based on associations between prognostic factor (depressive symptoms or depression) and health outcomes relevant to this systematic review

<sup>b</sup>High refers to univariate associations between prognostic factor (depressive symptoms or depression) and health outcomes relevant to this systematic review

Appendix XIII. List of possibly relevant articles published in non-English language (6 studies)

1. Heinrich M, Hafenbrack K, Michel C, Monstadt D, Marnitz U, Klinger R. Measures of success in treatment of chronic back pain: pain intensity, disability and functional capacity: determinants of treatment success in multimodal day clinic setting. Schmerz (Berlin, Germany). 2011 Jun 1;25(3):282-9.

• Sample size n=681

2. Mohr B, Graef T, Forster M, Krohn-Grimberghe B, Kurzeja R, Mantel F, Thomsen M, Hampel P. Influence of depressive symptoms and gender in chronic low back pain rehabilitation outcome: a pilot study. Die Rehabilitation. 2008 Oct 20;47(5):284-98.

• Sample size n=116

3. Pfingsten M, Hildebrandt J, Saur P, Franz C, Seeger D. Multidisciplinary treatment program on chronic low back pain, part 4. Prognosis of treatment outcome and final conclusions. Schmerz (Berlin, Germany). 1997 Feb 1;11(1):30-41.

• Sample size n=90

4. Roth KE, Kremer M, Maier GS, Sariyar M, Rompe JD, Kappis B. Epidural injection shows no advantages over oral medication and physiotherapy in the treatment of sciatica, irrespective of the duration of symptoms. Zeitschrift fur Orthopadie und Unfallchirurgie. 2014 Feb 27;152(1):46-52.

• Sample size n=106

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• Sample size n=177

6. Wu HF, Hsu TL, Hung SH, Tseng YL, Liu CL, Wang TJ. Preoperative disability and its influencing factors in patients with lumbar spondylolisthesis. Hu li za zhi The Journal of Nursing. 2018 Feb 1;65(1):33-41.

• Sample size n=86

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7. Truchon M, Côté D. Predictive validity of the Chronic Pain Coping Inventory in subacute low back pain. *Pain* 2005;116(3):205-12.

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10. Ritzwoller DP, Crounse L, Shetterly S, Rublee D. The association of comorbidities, utilization and costs for patients identified with low back pain. *BMC Musculoskelet Disord* 2006;7:72.

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