

# Depression Screening Tools for Patients with Kidney Failure

## A Systematic Review

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### Abstract

**Background and objectives** Patients with kidney failure experience depression at rates higher than the general population. Despite the Centers for Medicare and Medicaid Services' ESRD Quality Incentive Program requirements for routine depression screening for patients with kidney failure, no clear guidance exists.

**Design, setting, participants, & measurements** For this systematic review, we searched MEDLINE, PsycINFO, and other databases from inception to June 2020. Two investigators screened all abstracts and full text. We included studies assessing patients with kidney failure and compared a tool to a clinical interview or another validated tool (e.g., Beck Depression Inventory II). We abstracted data related to sensitivity and specificity, positive and negative predictive value, and the area under the curve. We evaluated the risk of bias using the Quality Assessment of Diagnostic Accuracy Studies 2.

**Results** A total of 16 studies evaluated the performance characteristics of depression assessment tools for patients with kidney failure. The Beck Depression Inventory II was by far the best studied. A wide range of thresholds were reported. Shorter tools in the public domain such as the Patient Health Questionnaire 9 and Geriatric Depression Scale 15 (adults over 60) performed well but were not well studied. Short tools such as the Beck Depression Inventory–Fast Screen may be a good option for an initial screen.

**Conclusions** There is limited research evaluating the diagnostic accuracy of most screening tools for depression in patients with kidney failure, and existing studies may not be generalizable to US populations. Studies suffer from limitations related to methodology quality and/or reporting. Future research should target widely used, free tools such as the Patient Health Questionnaire 2 and the Patient Health Questionnaire 9.

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### Introduction

The incidence and prevalence of kidney failure in the United States have increased steadily over the past 4 decades (1). Patients with kidney failure experience major depressive disorder at three to six times the rate of the general US population, depending on the method of assessment (2,3). Comorbid depression is associated with treatment nonadherence, poorer quality of life, worse sleep, more frequent emergency department visits, hospitalizations, suicide, and all-cause mortality (4–7).

The Centers for Medicare and Medicaid Services requires routine depression screening for patients with kidney failure as part of their ESRD Quality Incentive Program (ESRD-QIP) (8). However, there is no system-wide screening protocol, leading to wide variation in the way depression is assessed. Established evidence and guidelines suggest that screening for depression in the general population is both

accurate and can improve outcomes (9). However, screening may lead to false positives and concomitant iatrogenic harm from unnecessary pharmacotherapy or resource-heavy psychotherapy. Screening may also lead to false negatives in which depression goes untreated. Patients with kidney failure differ from the general population both because they experience higher rates of comorbid depression, and because they often have symptoms related to their underlying diagnoses and treatments that mimic the somatic symptoms of depression.

Given the wide variation in depression screening options and lack of a gold standard assessment tool for patients with kidney failure, a clear understanding of the validity of the available screening tools is vital. The purpose of this review is to identify depression screening tools (and/or thresholds) appropriate for patients with kidney failure, and to better understand the effect of depression screening in this population.

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## Materials and Methods

This is part of a larger systematic review commissioned by the Veterans Health Administration that examined both screening and the effectiveness of interventions for patients with kidney failure and comorbid depression (10). The protocol, which follows PRISMA guidelines (11), was registered to PROSPERO before study initiation (CRD42020140227).

### Data Sources and Searches

We searched Ovid MEDLINE, PsycINFO, Elsevier EM-BASE, and Ovid EBM Reviews Cochrane Database of Systematic Reviews (Database of Abstracts of Reviews of Effects, Health Technology Assessment Database, Cochrane CENTRAL, *etc.*) from database inception through June 2020. We reviewed the bibliographies of relevant articles and contacted experts to identify additional studies. Search strategies were developed in consultation with a research librarian, and were peer reviewed by a second research librarian using the instrument for Peer Review of Search Strategies (12; Supplemental Material). All studies identified were completed before the onset of the COVID-19 pandemic.

### Study Selection

Studies were eligible if they: (1) assessed depression in patients with kidney failure or stage 5 CKD; (2) compared an index (examined) tool to a “gold standard” clinical interview or another well-validated tool (*e.g.*, Beck Depression Inventory II [BDI-II] [13]); (3) were published in English; and (4) examined tools based on criteria from the Diagnostic and Statistical Manual of Mental Disorders IV (DSM-IV) or higher (Supplemental Table 1). Studies were independently reviewed by at least two reviewers. Discordant results were resolved through consensus or a third reviewer.

### Data Abstraction and Quality Assessment

From each study, we abstracted details related to sample size, setting, population, inclusion and exclusion criteria, administration and timing of depression screening, the index and reference standard (comparison), and findings. Data were abstracted by one investigator and confirmed by a second. Two reviewers independently assessed study risk of bias using the Quality Assessment of Diagnostic Accuracy Studies 2 (QUADAS-2). The QUADAS-2 is a commonly used 17-item tool for assessing the risk of bias across four domains: (1) flow and timing, (2) application of the reference standard, (3) application of the index test, and (4) patient selection (14). See Supplemental Table 2 for a list of all items. Disagreements were resolved by consensus or a third reviewer.

### Data Synthesis

We qualitatively synthesized findings, organized them in tables, and present forest plots of the summary measures (*e.g.*, sensitivity, specificity). The data did not allow for quantitative pooling of results.

## Results

We reviewed 8050 titles and abstracts and 189 full text studies. A total of 16 studies were included (Figure 1). Nine studies examined the performance of the BDI-II (13). Other tools include the Cognitive Depression Index (CDI) (15), the Center for Epidemiologic Studies–Depression Scale (CES-D) (16), the Hospital Anxiety and Depression Scale–Depressive Subscale (HADS-D) (17), the Geriatric Depression Scale 15 (GDS-15) (18,19), the Hamilton Depression Rating Scale (20), the Patient Health Questionnaire 9 (PHQ-9) (21), and others. We identified only one study of a depression tool specifically targeting patients on maintenance dialysis (Depression Inventory–Maintenance Hemodialysis [DI-MHD]) (22). Supplemental Table 3 and Table 1 provide study characteristics and Table 2 provides a brief description of the included tools and the gold standard interviews used as reference standards.

There were five US studies (24,33,35,37,46). Other studies were located in Australia (25), Canada (27), China (22), Italy (29), The Netherlands (30,36), Norway (34), Saudi Arabia (23), Turkey (28), and the United Kingdom (26,30).

Most studies included only patients undergoing hemodialysis. Only four studies also included participants undergoing peritoneal dialysis (32,34,35,37). Across studies reporting time on dialysis, the minimum (mean) months was 8.5 (interquartile range, 4–22) (34) and the maximum was 72.2 (SD=11.7) (28).

Of the 16 studies, 11 compared tools with a gold standard clinical interview (*e.g.*, Structured Clinical Interview for DSM-IV [38]), and five used other established, validated assessment measures (*e.g.*, BDI-II [13]) for comparison.

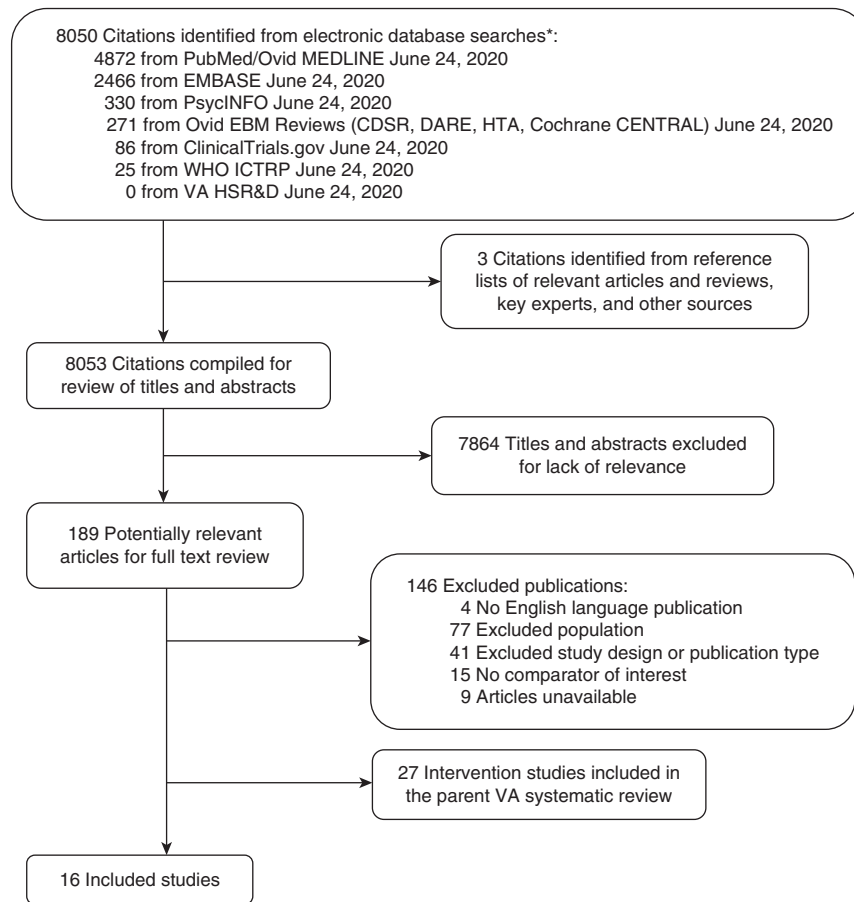
Seven studies examined thresholds for major depressive disorder (23–26,30,33,37), one of which also categorized less severe depression (23). The remaining nine studies did not describe differences between major depressive disorder, less severe depressive disorders (*e.g.*, dysthymia, pervasive depressive disorder), and subclinical symptoms (Supplemental Table 3) (22,27–29,31,32,34–36).

The 16 studies were relatively similar in quality, with the risk of bias largely unclear for patient selection, the index test, and the reference standard. Figure 2 summarizes the number of studies rated as low, unclear, and high risk of bias across the four QUADAS-2 domains (Supplemental Table 2 reports individual study ratings).

### Screening Tools Compared with a Gold Standard Clinical Interview

Included studies compared nine tools to clinical interviews, across a range of thresholds. Figure 3 illustrates the performance characteristics by tool and threshold. Supplemental Table 4 and Table 1 provide detail.

**Beck Depression Inventory II.** Five studies examined the accuracy of the BDI-II in diagnosing major depressive disorder compared with a gold standard clinical interview (24–26,30,37). Sample sizes ranged from 40 (26) to 96 (24). Two were conducted in the United States (24,37). One was a small, multicenter study ( $n=62$ ) that reported an optimal BDI-II cutoff of  $\geq 16$ . Sensitivity was 0.91 and specificity was 0.86, with an area under the curve (AUC) of 0.94 (37). The second was a multicenter study of adults 65 and older



**Figure 1. | Literature flow chart.** \*After deduplication. CDSR, Cochrane Database of Systematic Reviews; DARE, Database of Abstracts of Reviews of Effects; EBM, Evidence-based Medicine; HSR&D, Health Services Research and Development; HTA, Health Technology Assessment Database; ICTRP, International Clinical Trials Registry Platform; VA, Veterans Affairs; WHO, World Health Organization.

( $n=96$ ). At a cutoff of  $\geq 10$ , sensitivity was 0.68, specificity was 0.77, and reported AUC was 0.73 (24).

One study examined a range of thresholds (30). The optimal threshold was  $\geq 15$ , with a reported AUC of 0.93. Another study reported a much lower AUC (24). This study's population was limited to older adults, and age differences may have contributed to the difference in performance (24).

Among studies screening for depressive symptoms and disorders ranging from subclinical to major depressive disorder (22,31,32,34), sample sizes ranged from 43 (26) to 319 (22). Only one study ( $n=98$ ) was conducted in the United States (31). At a threshold of  $\geq 14$ , sensitivity was 0.62, specificity was 0.81, and AUC was 0.77 (31). The largest study ( $n=319$ ), conducted in China, compared the BDI-II ( $\geq 19$ ) to the Structured Clinical Interview for DSM-IV as part of a development and validation study for a depression screen designed specifically for patients undergoing maintenance hemodialysis (22). Sensitivity, specificity, positive predictive value, negative predictive value, and AUC were 0.83, 0.86, 0.63, 0.94, and 0.84, respectively.

**Cognitive Depression Index.** Four studies compared the CDI to a gold standard clinical interview (25,26,31,34), of which only two screened specifically for major depressive disorder (25,26). One study ( $n=45$ ) examined a threshold of

$\geq 11$ , with a sensitivity, specificity, and AUC of 0.79, 0.81, and 0.94, respectively (25). The second study identified an optimal threshold of  $\geq 10$ . Sensitivity, specificity, and AUC were 0.78, 0.81, and 0.94, respectively (26).

The two studies screening for the range of depressive symptoms and diagnoses examined thresholds of  $\geq 8$  ( $n=98$ ) (31) and  $\geq 11$  ( $n=109$ ) (34). Sensitivity was 0.50 (31) and 0.82 (34), specificity was 0.83 (31) and 0.93 (34), and AUC was 0.76 (31) and 0.89 (34).

Of note, two studies (26,34) compared the BDI-II and the CDI to a clinical interview and both concluded that the BDI-II performed better.

**Center for Epidemiologic Studies–Depression Scale.** A multisite study ( $n=98$ ) (31) compared the CES-D ( $\geq 18$ ) to the Structured Clinical Interview for DSM-IV for depressive disorders and subclinical symptoms. Sensitivity, specificity, and AUC were 0.69, 0.83, and 0.89, respectively.

**Depression Inventory–Maintenance Hemodialysis.** A single validation study ( $n=319$ ) conducted in China compared the Structured Clinical Interview for DSM-IV to both the BDI-II and the DI-MHD and found that at a cutoff of  $\geq 25$ , the DI-MHD performed better than the BDI-II. Sensitivity, specificity, and AUC were 0.95, 0.93, and 0.94, respectively (22).

Table 1. Study characteristics						
Author (Ref.)	N	Population	Index Test(s)	Reference Standard	% With Major Depressive Disorder Diagnosis <sup>a</sup>	
					Index Test (cutoff)	Reference Standard
Alsuwaida and Alwahhabi (23)	26	HD	SRQ (Arabic version)	Clinical interview	NR	15.4%
Balogun <i>et al.</i> (24)	96	Dialysis (HD/PD: NR)	BDI-II, GDS-15	Clinical interview	BDI-II ( $\geq 10$ ): 37.1%, GDS-15 ( $\geq 5$ ): 32.3% (subset $n=62$ )	30.6% (subset $n=62$ )
Bautovich <i>et al.</i> (25)	45	HD	BDI-II, CDI	Clinical interview	BDI-II/CDI: NR	13.3%
Chilcot <i>et al.</i> (26)	40	HD	BDI-II, CDI	MINI	BDI-II ( $\geq 16$ ) on dialysis: 32.5%, off dialysis: 30%, CDI ( $\geq 10$ ) on and off dialysis: 32.5%	22.5% (off dialysis)
Collister <i>et al.</i> (27) <sup>b</sup>	50	HD	Single question from the ESAS	HADS	ESAS: NR	HADS ( $\geq 7$ ): 54%
Gencoz <i>et al.</i> (28) <sup>b</sup>	45	HD	Ham-D	SCID-I (Turkish Translation)	Ham-D: NR	4% MDD (18% other depressive disorders)
Giordano <i>et al.</i> (29) <sup>b</sup>	31	HD	GDS-15	BDI-II	GDS-15 ( $\geq 6$ ): 58%	BDI-II ( $\geq 14$ ): 61%
Grant <i>et al.</i> (30) <sup>c</sup>	57	HD	BDI-II	Clinical interview (based on ICD-10 diagnosis)	BDI-II ( $\geq 10$ ): 56.1%, BDI-II ( $\geq 15$ ): 31.6%	12.3%
Hedayati <i>et al.</i> (31) <sup>b</sup>	98	HD	BDI-II, CDI, CES-D, Feinstein Scale	SCID-I	BDI-II ( $\geq 14$ ): 30.6%, CES-D ( $\geq 18$ ): 30.6%	17.3% MDD (26.5% any depressive disorders)
Loosman <i>et al.</i> (32) <sup>b</sup>	62	HD and PD	BDI-II, HADS	MINI	BDI-II, HADS: NR; 33.9%	
Neitzer, 2012 (33)	134	HD	BDI-FS	BDI-II	BDI-FS ( $\geq 4$ ): 30.1%	BDI-II ( $\geq 16$ ): 28.7%
Preljevic <i>et al.</i> (34) <sup>b</sup>	109	HD and PD	BDI-II, CDI, HADS-D	SCID-I	BDI-II ( $\geq 16$ ): 20.8%, CDI ( $\geq 11$ ): NR, HADS-D ( $\geq 8$ ): 20.1%	14.7% MDD (22% any depressive disorders)
Troidle <i>et al.</i> (35) <sup>b</sup>	97	CPD and HD	Two items from the KDQOL SF-36	BDI-II	KDQOL SF-36: NR	BDI-II ( $\geq 11$ ): NR
van den Beukel <i>et al.</i> (36) <sup>b</sup>	133	HD: 72%	MHI5 of the SF-36	BDI-II/CDI (Dutch Translation)	MHI5 ( $\leq 70$ ): 39%	BDI-II ( $\geq 16$ ): 23% CDI ( $\geq 10$ ): 23%
Watnick <i>et al.</i> (37)	62	HD and PD	BDI-II, PHQ-9	SCID-I	BDI-II, PHQ-9: NR	19.4%
Wang <i>et al.</i> (22) <sup>b</sup>	319	HD	BDI-II, DI-MHD (Chinese language)	SCID-I	BDI-II ( $\geq 19$ ): 20.7%, DI-MHD ( $\geq 25$ ): 20%	21.9%

HD, hemodialysis; SRQ, Self-Reporting Questionnaire; NR, not reported; PD, peritoneal dialysis; BDI-II, Beck Depression Inventory II; GDS-15, Geriatric Depression Scale 15; CDI, Cognitive Depression Index; MINI, Mini International Neuropsychiatric Interview; ESAS, Edmonton Symptom Assessment System; HADS: Hospital Anxiety and Depression Scale; Ham-D, Hamilton Depression Rating Scale; SCID-I, Structured Clinical Interview for Diagnostic and Statistical Manual of Mental Disorders IV; MDD, major depressive disorder; ICD-10, 10th revision of the International Statistical Classification of Diseases and Related Health Problems; CES-D, Center for Epidemiologic Studies–Depression Scale; BDI-FS, Beck Depression Inventory–Fast Screen; HADS-D, Hospital Anxiety and Depression Scale–Depressive Subscale; CPD, chronic pulmonary disease; KDQOL SF-36, Kidney Disease Quality of Life Short Form 36; MHI5, Mental Health Inventory 5; PHQ-9, Patient Health Questionnaire 9; DI-MHD, Depression Inventory–Maintenance Hemodialysis.

<sup>a</sup>The reported prevalence of major depressive disorder identified by the index test and the reference standard.

<sup>b</sup>Screened for depressive symptoms or milder forms of depression in addition to major depressive disorder.

<sup>c</sup>Included cutoff values for both major depressive disorder and for milder forms of depression and subclinical symptoms.

**Table 2. Characteristics of included screening tools and gold standard semi-structured diagnostic interviews examined**

Name	Abbrev.	Number of Items	Description	Time to Complete/Score
<b>Gold standard semi-structured diagnostic interviews</b>				
Mini International Neuropsychiatric Interview	MINI	NA	A short, structured diagnostic interview, developed jointly by psychiatrists and clinicians in the United States and Europe, for DSM-IV and ICD-10 psychiatric disorders.	Administration: 15 min
Structured Clinical Interview for DSM-IV Axis I Disorders (38)	SCID-I	NA	A semi-structured interview administered by a trained mental health professional familiar with DSM-IV Axis I diagnostic criteria.	Administration: 1–2 h
<b>Screening tools</b>				
Beck Depression Inventory II (13,39)	BDI-II	21	Widely used, validated self-report tool designed to assess depression severity in adolescents and adults. Most widely studied instrument in the kidney failure population. Closely mirrors DSM-IV criteria for major depressive disorder, and includes questions related to cognitive, affective, and somatic symptoms.	Respondent: 5–10 min Administrator: 5 min
Beck Depression Inventory II Fast Screen (40)	BDI-FS	7	Brief version of the BDI-II that excludes somatic symptoms. Designed to screen for major depressive disorder in medical patients.	Respondent: <5 min Administrator: <3 min, can be staff
Cognitive Depression Index (41)	CDI	15	A subset of the BDI-II (first 15 items), eliminating items related to somatic symptoms. It was developed for use in patients with CKD, with the goal of reducing the likelihood of the overdiagnosis of depression.	Respondent: 7–8 min Administrator: 5 min. These are estimates based on the BDI-II from which it is derived.
Center for Epidemiologic Studies–Depression Scale (16,39)	CES-D	20	Widely used, revised in 2004, evaluates depressive symptoms across four factors: depressive affect, well-being, somatic symptoms, and interpersonal relations.	Respondent: <10 min Administrator: <10 min (can be scored during administration)
Depression Inventory–Maintenance Hemodialysis (22)	DI-MHD		Developed specifically for patients with kidney failure.	Respondent: 5 min Administrator: no information yet available.
Edmonton Symptom Assessment System (27)	ESAS <sup>a</sup>	1	Single item that asks patients to rate depression from 0 (not depressed) to 10 (worst depression) (1).	Respondent: minimal Administrator: minimal
Hamilton Depression Rating Scale (20,42)	Ham-D	17	Assesses the frequency and intensity of depressive symptoms. Developed in 1960, and last revised in 1967 (20).	Administrator: 15–20 min
Hospital Anxiety and Depression Scale–Depression Subscale (17)	HADS-D	7	Subscale of the 14-item HADS that includes ratings of physical, cognitive, and affective symptoms of depression (17).	Respondent: ≤5 min (for entire HADS including anxiety items) Administrator: 1–2 min
Geriatric Depression Scale-15 (18,39)	GDS-15	15	Shortened version of the original 30-item GDS, assesses depressive symptoms in older adults, developed in 1982 (18,19).	Respondent: 2–5 min Administrator: 2 min
Kidney Disease Quality of Life Short Form - 36 (43)	KDQOL SF-36	1	The KDQOL SF-36 is a self-report measure developed for patients with kidney disease on dialysis. Six-options ranging from “all of the time” to “none of the time.” The included study tests the use of two single questions. (1) “Have you felt so down in the dumps that nothing could cheer you up?” (2) “Have you felt downhearted and blue?”	Respondent: minimal Administrator: minimal
Mental Health Inventory 5 (44)	MHI5	5	Also known as the mental health subscale of the SF-36. Six option Likert scale ranging from “all of the time” to “none of the time.”	Respondent: <5 min Administrator: minimal
Patient Health Questionnaire-9 (21,39)	PHQ-9	9	Developed in 2001. Screen of depression and severity. Widely used in the United States and internationally (21).	Respondent: <3 min Administrator: minimal

Name	Abbrev.	Number of Items	Description	Time to Complete/Score
Self-Reporting Questionnaire (45)	SRQ	20	Developed by the WHO to screen for a range of mental health disorders (45).	Respondent: "a few min" Administrator: 1 min, yes/no additive

MINI, Mini International Neuropsychiatric Interview; NA, not applicable; DSM-IV, Diagnostic and Statistical Manual of Mental Disorders IV; ICD-10, 10th revision of the International Statistical Classification of Diseases and Related Health Problems; SCID-I, Structured Clinical Interview for DSM-IV; BDI-II, Beck Depression Inventory–II; BDI-FS, Beck Depression Inventory–Fast Screen; CDI, Cognitive Depression Index; CES-D, Center for Epidemiologic Studies–Depression Scale; DI-MHD, Depression Inventory–Maintenance Hemodialysis; ESAS, Edmonton Symptom Assessment System; Ham-D, Hamilton Depression Rating Scale; HADS-D, Hospital Anxiety and Depression Scale–Depressive Subscale; GDS-15, Geriatric Depression Scale-15; KDQOL SF-36, Kidney Disease Quality of Life Short Form 36; SF-36, 36-item Short-Form Health Survey Questionnaire; MHI5, Mental Health Inventory 5; PHQ-9, Patient Health Questionnaire 9; SRQ, Self-Reporting Questionnaire; WHO, World Health Organization.

<sup>a</sup>The ESAS is a ten-item scale. However, only one item applies to depression.

**Hamilton Depression Rating Scale** A single study ( $n=45$ ) conducted in Turkey compared the Hamilton Depression Rating Scale ( $\geq 10$ ) to the Structured Clinical Interview for DSM-IV and screened for the range of depressive symptoms and disorders. Reported sensitivity was 1.00, specificity was 0.80, and AUC was 0.85 (28).

**Hospital Anxiety and Depression Scale–Depression Subscale.** Two studies examined the performance characteristics of the HADS-D (32,34). Both studies screened for depressive disorders and subclinical symptoms. One study ( $n=62$ ;  $\geq 6$ ) reported sensitivity, specificity, and AUC values of 0.91, 0.76, and 0.89, respectively (32). The other ( $n=109$ ;  $\geq 8$ ) reported sensitivity, specificity, and AUC values of 0.73, 0.87, and 0.91, respectively. Of note, this study also examined the BDI-II ( $\geq 16$ ), and concluded it performed better than the HADS-D (34).

**Geriatric Depression Scale 15 (adults aged 60+).** A single study ( $n=96$ ) compared the GDS-15 ( $\geq 5$ ) to a gold standard interview for major depressive disorder. Sensitivity was 0.62, specificity was 0.82, and AUC was 0.81 (24).

**Patient Health Questionnaire 9.** A small multisite study ( $n=62$ ) compared the PHQ-9 ( $\geq 10$ ) to the Structured Clinical Interview for DSM-IV for major depressive disorder. Sensitivity and specificity were both 0.92, and AUC was 0.94 (37).

**Self-Reporting Questionnaire.** A single small study ( $n=26$ ) conducted in Saudi Arabia compared the Self-Reporting Questionnaire ( $\geq 13$ ) to Structured Clinical Interview for DSM-IV for major depressive disorder.

Sensitivity, specificity, and AUC were 1.00, 0.82, and 0.96, respectively (23).

### Screening Tools Compared with Other Tools

Five studies compared other, generally short, tools to established, validated tools (*i.e.*, BDI-II, HADS-D; see Figure 4, Supplemental Table 5, Table 1) (27,29,33,35,36). One study screened for major depressive disorder specifically, and evaluated the BDI–Fast Screen (40). It had high sensitivity and specificity compared with the BDI-II ( $\geq 16$ ) (33). The GDS-15 ( $\geq 6$ ) screened for a range of depression diagnoses and symptoms and appeared to perform well (29).

### Timing of Screening

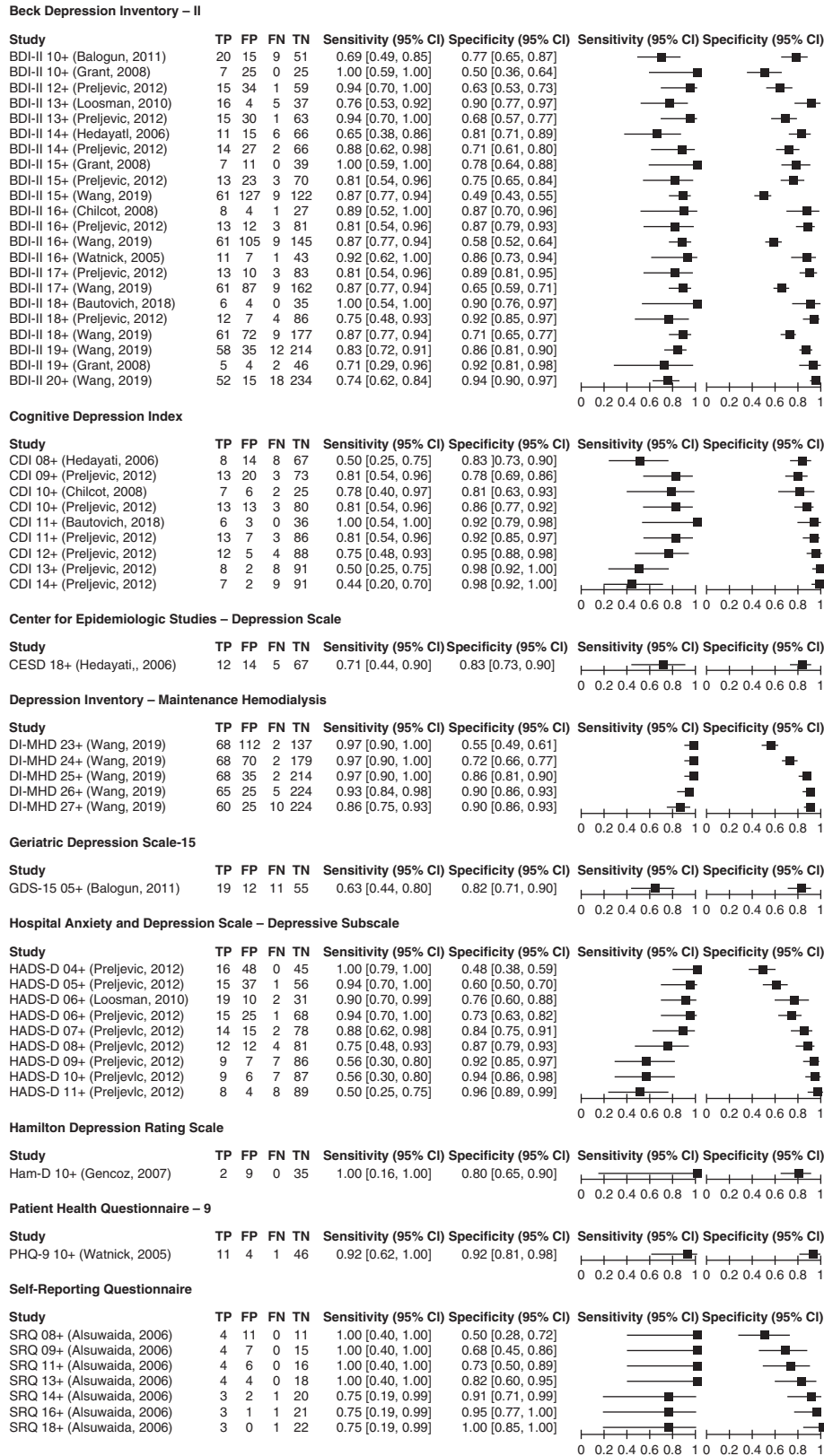
A small ( $n=43$ ) multisite UK study in outpatient hemodialysis units compared depression screening (BDI-II, CDI) completed on and off dialysis (26). Findings indicated a high level of agreement among patients who were depressed. However, patients who were not depressed had higher mean overall BDI-II (9.6 [6.2] versus 7.3 [5.7],  $P=0.007$ ) and somatic symptom item scores (4.4 [2.5] versus 3.3 [2.1],  $P=0.01$ ) on assessments completed while undergoing dialysis (Supplemental Table 3).

### Discussion

We identified 16 studies examining the performance characteristics of depression screening tools in patients



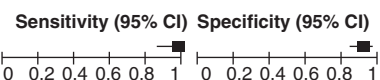
**Figure 2. | QUADAS-2 risk of bias summary.** Quality Assessment of Diagnostic Accuracy Studies 2 (QUADAS-2), independently assessed by two investigators.



**Figure 3. | Performance characteristics of tools compared with a gold standard clinical interview.** BDI-II, Beck Depression Inventory-II; CES-D, Center for Epidemiologic Studies-Depression Scale; CI, confidence interval; DI-HMD, Depression Inventory-Maintenance Hemodialysis; FN, false negative; FP, false positive; GDS-15, Geriatric Depression Scale-15; HADS-D, Hospital Anxiety and Depression Scale-Depressive Subscale; Ham-D, Hamilton Depression Rating Scale; SRQ, Self-Reporting Questionnaire; TN, true negative; TP, true positive.

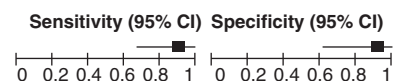
## Beck Depression Inventory – Fast Screen

Study	TP	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)
BDI-FS 4+ (Neltzer, 2012)	37	8	1	88	0.97 [0.86, 1.00]	0.92 [0.84, 0.96]



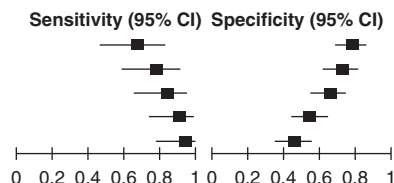
## Geriatric Depression Scale – 15

Study	TP	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)
GDS-15 6+ (Giordano, 2007)	17	1	2	11	0.89 [0.67, 0.99]	0.92 [0.62, 1.00]



## Mental Health Inventory 5

Study	TP	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)
MHI5 66+ (Van den Beukel, 2012)	20	23	10	80	0.67 [0.47, 0.83]	0.78 [0.68, 0.85]
MHI5 70+ (Van den Beukel, 2012)	24	29	7	74	0.77 [0.59, 0.90]	0.72 [0.62, 0.80]
MHI5 74+ (Van den Beukel, 2012)	25	36	5	67	0.83 [0.65, 0.94]	0.65 [0.55, 0.74]
MHI5 78+ (Van den Beukel, 2012)	28	47	3	55	0.90 [0.74, 0.98]	0.54 [0.44, 0.64]
MHI5 82+ (Van den Beukel, 2012)	28	56	2	46	0.93 [0.78, 0.99]	0.45 [0.35, 0.55]



**Figure 4. | Performance characteristics of tools compared with a gold standard clinical interview.** BDI-FS, Beck Depression Inventory–Fast Screen; ESAS, Edmonton Symptom Assessment System; MHI5, Mental Health Inventory 5.

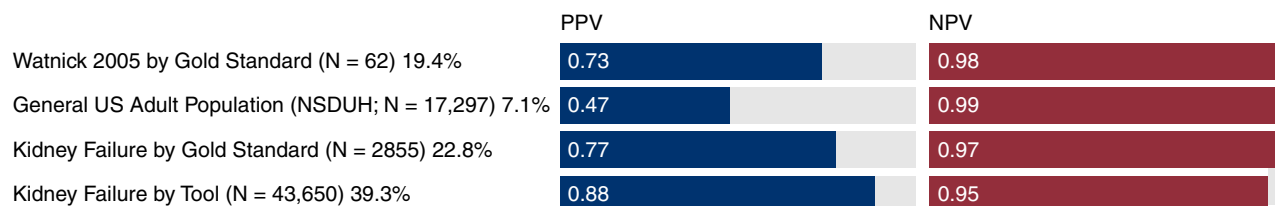
with kidney failure, and found depression can be accurately diagnosed. By far, the strongest body of evidence addressed the use of longer screening tools and those that require a per-administration cost (*e.g.*, BDI-II), which are not common in medical settings. We found promising evidence that shorter instruments in the public domain such as the PHQ-9 and GDS-15 (for older adults) performed well, but only one study compared each of these instruments to a clinical interview (24,37).

Across studies, sample sizes were small, and studies examined a wide (and inconsistent) range of thresholds. In addition, methodologic details, particularly related to the selection of patients and the conduct and/or interpretation of both the index and reference tests, were generally poorly reported. We identified few studies conducted in the United States, or countries with similar health systems, raising concerns about the generalizability of findings. Except for the BDI-II, the evidence base is quite limited due to few studies examining each tool. There was heterogeneity in how depression was operationalized across studies. Half of the studies evaluated the performance characteristics associated with thresholds intended to screen for major depressive disorder, whereas the other half defined depression broadly, some including subclinical depressive symptoms.

Figure 5 illustrates the effect of sensitivity and specificity across different population-based depression rates. We

used data from a US study comparing the PHQ-9 to a gold standard interview (37) that screened specifically for major depressive disorder. At a threshold of  $\geq 10$ , both sensitivity and specificity were 0.92. Holding these constant, we compared positive and negative predictive values across reported major depressive disorder prevalence rates for (1) general US populations (7.1%) (2); (2) US patients with kidney failure, diagnosed using a gold standard interview (22.8%) (3); and (3) US patients with kidney failure, diagnosed using a screening tool (39.3%) (3). Across populations, the negative predictive values, or accuracy of eliminating depression, are generally high, and false negatives are unlikely. However, the positive predictive values, or accuracy of correctly diagnosing depression, range from 0.47 to 0.88, suggesting in this example that for populations with a lower prevalence of depression, the potential for false positives may be high (Figure 5). Providers should keep these factors in mind when using the results of depression screening tools to guide treatment decisions.

Among the studies evaluating the BDI-II as a tool to identify major depressive disorder, the threshold that best optimized the balance between sensitivity and specificity for patients with kidney failure was  $\geq 16$ . In fact, in some studies, the BDI-II performed reasonably well when compared with a gold standard clinical interview. The caveats



**Figure 5. | PHQ-9  $\geq 10$  positive and negative predictive values: Three US subpopulations.** Both sensitivity and specificity are held constant at 0.92 based on findings from Watnick's 2005 study comparing the PHQ-9 to a gold standard clinical interview (37). Prevalence data for the general US population comes from the 2017 National Survey on Drug Use and Health (2), and prevalence for patients with kidney disease is reported in a meta-analysis of 249 unique populations (3). NPV, negative predictive value; PHQ-9, Patient Health Questionnaire 9; PPV, positive predictive value.



are the heterogeneity in how tools were administered, and that very few studies contributed data for the same thresholds. Interestingly, two studies found that, compared with a clinical interview, the BDI-II performed better than the CDI, a subset of the BDI without items related to somatic symptoms.

Shorter screening tools compared with established, validated tools performed well overall. Because the ESRD-QIP requires a follow-up after an initial positive screen, these short tools may be appropriate for an initial depression screen of all patients with kidney failure. In particular, the BDI-Fast Screen performed well when compared with the BDI-II. Of note, we identified no studies evaluating the PHQ-2, arguably the most commonly used short screen for depression in US medical settings.

One study examined differences in performance based on the timing of screening and found that participants who were not depressed reported significantly more somatic symptoms when they were screened during dialysis sessions versus off dialysis. Not only were scores on somatic items significantly higher, but BDI-II scores were significantly higher as well. This has implications for dialysis units working to streamline processes, as it illustrates the potential for overdiagnosis and overtreatment.

There are several important limitations; notably, small sample sizes and few studies examining specific tool thresholds. Many studies were conducted outside of the United States and examined participants and health systems that differ from US populations and settings. In addition, the lack of methodology detail reported in many of the studies contributed to uncertainty about study processes and poor or unclear quality ratings. The definition of depression varied widely, which hampered our ability to synthesize the body of research for each tool. Future studies should use standardized language and diagnostic criteria (*e.g.*, DSM-5) (47).

As described above, future research examining the diagnostic accuracy of depression tools in US populations is needed. In addition, although the PHQ-9 is used widely in medical settings, current research in kidney failure populations is extremely limited. Similarly, despite wide use of the PHQ-2 as an initial screen, no studies were identified. Research evaluating performance characteristics of both tools in this population are warranted. There are a handful of studies supporting the use of the BDI-II as a screening tool for major depressive disorder in this population. However, the BDI-II requires a per-use fee, is more commonly used in research and mental health than in medical settings, was developed to align to the out-of-date DSM-IV, and may be less informative for screening and assessment due to its reliance on somatic symptoms. Future research should target free tools (*e.g.*, CES-D, PHQ-9, PHQ-2) that are widely used in US medical settings.

Short, population-targeted tools may be appropriate as an initial screen for depression in dialysis settings, and we identified several high-performing tools that used the BDI-II as a reference standard. However, more research is needed to validate existing findings. Given the ESRD-QIP requirement of both an initial and follow-up screen (if warranted), future research should evaluate both quick screens and those that are more comprehensive. Finally, the DI-MHD appears to be the only screening tool for

depression designed specifically for patients with kidney failure. It performed well in a large sample in China. Additional research validating the DI-MHD generally and in English-speaking patients has the potential to affect screening practices in this population.

We identified no studies examining the effect of screening on health outcomes, and only one study that examined differences in implementation. It compared differences in both overall BDI-II scores and somatic item scores when completed on versus off dialysis and touches on only one of many important depression screening implementation issues (*e.g.*, timing, location, administration). Implementation is an important issue, as patients' responses may differ based on setting and timing, due not only to the experience of somatic symptoms, but also the perception of privacy and other factors. Future implementation research may help to better identify depression in patients with kidney failure and minimize overtreatment. We identified no studies examining potential harm associated with the lack of a standardized depression screening tool (*e.g.*, false positive or negative depression screens contributing to over- or undertreatment). Harm research is especially important in patients with chronic conditions such as kidney failure, given that the physical symptoms of chronic illnesses and their treatment can mimic the somatic symptoms of depression. Also important, but missing, is evidence of potential demographic and clinical differences. Research in these areas will help decision makers to implement screening processes that are not only evidence based, but also the best fit for patient populations.

Our findings have implications for the selection and implementation of depression screening in patients with kidney failure, and highlight the moderate positive predictive values in this population. Clinicians should be prepared to validate positive screens before making treatment decisions that may be burdensome or introduce the possibility of harm.

There is limited research evaluating the diagnostic accuracy of most screening tools for depression in patients with kidney failure, and existing studies may not be generalizable to US populations. Studies suffer from limitations related to methodological quality and/or reporting. Future research should target widely used, free tools such as the PHQ-2 and the PHQ-9.

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### Supplemental Material

This article contains the following supplemental material online at <http://cjasn.asnjournals.org/lookup/suppl/doi:10.2215/CJN.05540420/-/DCSupplemental>.

Supplemental Material. Search strategies (parent VA systematic review) and supplemental references.

Supplemental Table 1. PICOTS by key question.

Supplemental Table 2. QUADAS-2 risk of bias assessment.

Supplemental Table 3. Characteristics of studies examining the diagnostic accuracy of depression screening tools in patients with kidney failure.

Supplemental Table 4. Findings of studies examining the diagnostic accuracy of depression screening tools in patients with kidney failure compared with a gold standard diagnostic interview.

Supplemental Table 5. Studies comparing a depression tool to another validated depression tool.

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See related Patient Voice, “Depression: A Side Effect of CKD,” on pages xxx–xxx and editorial, “Screening for Depression in People with Kidney Failure,” on pages xxx–xxx, respectively.

## **Depression Screening Tools for Patients with Kidney Failure: A Systematic Review Supplementary Material**

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## Supplemental Material. Search Strategies (Parent VA systematic review)

### Ovid MEDLINE(R) and Epub Ahead of Print, In-Process & Other Non-Indexed Citations and Daily 1946 to June 24, 2020

#	Searches
1	Kidney Failure, Chronic/
2	((((chronic or endstage or end-stage or endstate or end-state or failure or long-term or maintenance) adj2 (kidney or renal)) or ESKD or ESKF or ESRD or ESRF).ti,ab,kf.
3	Renal Dialysis/ or Hemofiltration/ or Hemodiafiltration/ or Hemodialysis, Home/ or Peritoneal Dialysis/ or Peritoneal Dialysis, Continuous Ambulatory/
4	(dialysis or haemodiafiltration or hemodiafiltration or haemo-diafiltration or hemo-diafiltration or haemofiltration or hemofiltration or haemo-filtration or hemo-filtration or haemodialysis or hemodialysis or haemo-dialysis or hemo-dialysis).ti,ab,kf.
5	or/1-4
6	Depression/ or Depressive Disorder/ or Depressive Disorder, Major/ or Depressive Disorder, Treatment-Resistant/
7	(depressed or depressive or depression* or suicidal or suicide or suicides).ti,ab,kf.
8	or/6-7
9	and/5,8
10	Brief Psychiatric Rating Scale/ or Diagnostic Self Evaluation/ or "Diagnostic Techniques and Procedures"/ or Mental Status Schedule/ or Neuropsychological Tests/ or Patient Health Questionnaire/ or Psychiatric Status Rating Scales/ or exp Psychological Tests/ or exp "Surveys and Questionnaires"/
11	(checklist* or check-list* or questionnaire or questionnaires or instrument or instruments or inventory or inventories or scale or scales or schedule or schedules or screen or screened or screening or "Beck Depression" or BDI or BDI2 or "geriatric depression scale" or GDS or "Hamilton Rating Scale for Depression" or "Hospital Anxiety and Depression Scale" or HADS or "Kidney Disease Quality of Life" or KDQOL or "Medical Outcomes Study Short Form Health Survey 36" or MOS-SF36 or "Minnesota Multiphasic Personality Inventory" or MMPI or "Multiple Affect Adjective Check List" or MAACL or "Patient Health Questionnaire" or PHQ2 or PHQ-2 or PHQ9 or PHQ-9 or "PRIME-MD" or "Epidemiologic Studies Depression Scale" or CED or CESD or BREF or DASS21 or IDID or "Quick Inventory of Depressive Symptomatology Self-Report" or QIDS-SR or "RAND 36-Item Health Survey" or RAND-36 or "short form 36" or SF-36 or "Structured Clinical Interview" or SCID or "self-rating depression scale" or SDS or "Short Form Health Survey 36" or SF36).ti,ab,kf.
12	exp Behavior Therapy/ or exp Cognitive Behavioral Therapy/ or exp Mental Health Services/ or exp Psychotherapy/ or Psychosocial Support Systems/ or Social Support/ or Motivational Interviewing/ or Patient Participation/
13	(cognitive-behavior* or cognitive-behaviour* or intervention or interventions or nondrug or non-drug or nonpharmac* or non-pharmac* or pharmac* or program* or psych* or psychosocial or psycho-social or rehabilitation or therapy or therapies or treat*).ti,ab,kf.
14	Antidepressive Agents/ or Antidepressive Agents, Second-generation/ or Serotonin Uptake Inhibitors/ or "Serotonin and Noradrenaline Reuptake Inhibitors"/ or Adrenergic Uptake Inhibitors/ or 5-hydroxytryptophan/ or Amisulpride/ or Bupropion/ or Citalopram/ or Fluoxetine/ or Fluvoxamine/ or Maprotiline/ or Mianserin/ or Paroxetine/ or Quipazine/ or Ritanserin/ or Sulpiride/ or Trazodone/ or Tryptophan/ or Venlafaxine Hydrochloride/ or Viloxazine/
15	(antidepress* or anti-depress* or 5-hydroxytryptophan or Amisulpride or Bupropion or Citalopram or Escitalopram or Fluoxetine or Fluvoxamine or Maprotiline or Mianserin or Paroxetine or Quipazine or Ritanserin or Sulpiride or Trazodone or Tryptophan or Venlafaxine Hydrochloride or Viloxazine or SSRI or SSRIs or "selective serotonin reuptake" or "selective serotonin re-uptake" or SNRI or SNRIs or "Serotonin and Noradrenaline Reuptake Inhibitor" or "Serotonin and Noradrenaline Reuptake Inhibitors" or NRI or NRIs or "norepinephrine reuptake inhibitor" or "norepinephrine reuptake inhibitors").ti,ab,kf.

16	Antidepressive Agents, Tricyclic/ or Amitriptyline/ or Amoxapine/ or Clomipramine/ or Desipramine/ or Dothiepin/ or Doxepin/ or Imipramine/ or Iprindole/ or Lofepramine/ or Nortriptyline/ or Opipramol/ or Protriptyline/ or Trimipramine/
17	(Amitriptyline or Amoxapine or Clomipramine or Desipramine or Dothiepin or Doxepin or Imipramine or Iprindole or Lofepramine or Nortriptyline or Opipramol or Protriptyline or Trimipramine or Gabapentin or Sildenafil or Vardenafil).ti,ab,kf,nm.
18	(nondrug or non-drug or nonpharmacolog* or non-pharmacolog*).ti,ab,kf.
19	Exercise Therapy/ or Resistance Training/
20	((((aerobic or resistance) adj2 (exercis* or program* or therap* or train*)) or (exercise adj2 (program* or therap* or train*)) or cross-training).ti,ab,kf.
21	Music Therapy/
22	music therapy.ti,ab,kf.
23	or/10-22
24	and/9,23
25	limit 24 to english language

### PsycINFO 1806 to week of June 24, 2020

	Searches
1	Kidney Diseases/
2	((((chronic or endstage or end-stage or endstate or end-state or failure or long-term or maintenance) adj2 (kidney or renal)) or ESKD or ESKF or ESRD or ESRF).ti,ab.
3	Dialysis/ or Hemodialysis/
4	(dialysis or haemodiafiltration or hemodiafiltration or haemo-diafiltration or hemo-diafiltration or haemofiltration or hemofiltration or haemo-filtration or hemo-filtration or haemodialysis or hemodialysis or haemo-dialysis or hemo-dialysis).ti,ab.
5	or/1-4
6	"Depression (Emotion)"/ or Major Depression/ or Reactive Depression/ or Recurrent Depression/ or Treatment Resistant Depression/
7	(depressed or depressive or depression* or suicidal or suicide or suicides).ti,ab.
8	or/6-7
9	and/5,8
10	exp Measurement/ or exp Attitude Measures/ or exp "Checklist (Testing)"/ or exp Inventories/ or exp Psychological Assessment/ or exp Questionnaires/ or exp Rating Scales/ or exp Screening/ or exp Screening Tests/ or exp Standardized Tests/ or exp "Stress and Coping Measures"/ or exp Testing/
11	(checklist* or check-list* or questionnaire or questionnaires or instrument or instruments or inventory or inventories or scale or scales or schedule or schedules or screen or screened or screening or "Beck Depression" or BDI or BDI2 or "geriatric depression scale" or GDS or "Hamilton Rating Scale for Depression" or "Hospital Anxiety and Depression Scale" or HADS or "Kidney Disease Quality of Life" or KDQOL or "Medical Outcomes Study Short Form Health Survey 36" or MOS-SF36 or "Minnesota Multiphasic Personality Inventory" or MMPI or "Multiple Affect Adjective Check List" or MAACL or "Patient Health Questionnaire" or PHQ2 or PHQ-2 or PHQ9 or PHQ-9 or "PRIME-MD" or "Epidemiologic Studies Depression Scale" or CED or CESD or BREF or DASS21 or IDID or "Quick Inventory of Depressive Symptomatology Self-Report" or QIDS-SR or "RAND 36-Item Health Survey" or RAND-36 or "short form 36" or SF-36 or "Structured Clinical Interview" or SCID or "self-rating depression scale" or SDS or "Short Form Health Survey 36" or SF36).ti,ab.
12	("Beck Depression" or BDI or BDI2 or "geriatric depression scale" or GDS or "Hamilton Rating Scale for Depression" or "Hospital Anxiety and Depression Scale" or HADS or "Kidney Disease Quality of Life" or KDQOL or "Medical Outcomes Study Short Form Health Survey 36" or MOS-SF36 or "Minnesota Multiphasic Personality Inventory" or MMPI or "Multiple Affect Adjective Check List" or MAACL or "Patient Health

	Questionnaire" or PHQ2 or PHQ-2 or PHQ9 or PHQ-9 or PHQ-ADS or "PRIME-MD" or "Epidemiologic Studies Depression Scale" or CED or CESD or BREF or DASS21 or IDID or "Quick Inventory of Depressive Symptomatology Self-Report" or QIDS-SR or "RAND 36-Item Health Survey" or RAND-36 or "short form 36" or SF-36 or "Structured Clinical Interview" or SCID or "self-rating depression scale" or SDS or "Short Form Health Survey 36" or SF36).tm.
13	exp Treatment/
14	exp Behavior Modification/ or exp Behavior Therapy/ or Biofeedback Training/ or Contingency Management/ or Self-management/ or Anxiety Management/ or Cognitive Therapy/ or Readiness to Change/ or Relaxation Therapy/ or Self-help Techniques/ or Self-monitoring/ or Stress Management/
15	(cognitive-behavior* or cognitive-behaviour* or intervention or interventions or nondrug or non-drug or nonpharmac* or non-pharmac* or pharmac* or program* or psych* or psychosocial or psycho-social or rehabilitation or therapy or therapies or treat*).ti,ab.
16	Antidepressant Drugs/ or Serotonin Norepinephrine Reuptake Inhibitors/ or Serotonin Reuptake Inhibitors/ or Tricyclic Antidepressant Drugs/ or Bupropion/ or Citalopram/ or Fluoxetine/ or Fluvoxamine/ or "Hydroxytryptophan (5-)"/ or MAPROTILINE/ or Mianserin/ or Paroxetine/ or RITANSERIN/ or Sulpiride/ or Trazodone/ or Tryptophan/ or Venlafaxine/
17	(antidepress* or anti-depress* or 5-hydroxytryptophan or Amisulpride or Bupropion or Citalopram or Escitalopram or Fluoxetine or Fluvoxamine or Maprotiline or Mianserin or Paroxetine or Quipazine or Ritanserin or Sulpiride or Trazodone or Tryptophan or Venlafaxine Hydrochloride or Viloxazine or SSRI or SSRIs or "selective serotonin reuptake" or "selective serotonin re-uptake" or SNRI or SNRIs or "Serotonin and Noradrenaline Reuptake Inhibitor" or "Serotonin and Noradrenaline Reuptake Inhibitors" or NRI or NRIs or "norepinephrine reuptake inhibitor" or "norepinephrine reuptake inhibitors").ti,ab.
18	(Amitriptyline or Amoxapine or Clomipramine or Desipramine or Dothiepin or Doxepin or Imipramine or Iprindole or Lofepramine or Nortriptyline or Opipramol or Protriptyline or Trimipramine or Gabapentin or Sildenafil or Vardenafil).ti,ab.
19	(nondrug or non-drug or nonpharmacolog* or non-pharmacolog* or coping or psychosocial* or psycho-social* or "social support" or "social work*" or stress).ti,ab.
20	exp Exercise/
21	((((aerobic or resistance) adj2 (exercis* or program* or therap* or train*)) or (exercise adj2 (program* or therap* or train*)) or cross-training).ti,ab.
22	Music Therapy/
23	music therapy.ti,ab.
24	or/10-23
25	and/9,24
26	limit 25 to english language

## Embase.com

Date searched: **June 24, 2020**

#	Search
#26	#25 AND [embase]/lim NOT ([embase]/lim AND [medline]/lim)
#25	#9 AND #23 AND [english]/lim
#24	#9 AND #23
#23	#10 OR #11 OR #12 OR #13 OR #14 OR #15 OR #16 OR #17 OR #18 OR #19 OR #20 OR #21 OR #22
#22	'music therapy':ti,ab,kw
#21	'music therapy'/de
#20	((((aerobic OR resistance) NEAR/2 (exercis* OR program* OR therap* OR train*)):ti,ab,kw) OR ((exercise NEAR/2 (program* OR therap* OR train*)):ti,ab,kw) OR 'cross training':ti,ab,kw
#19	'kinesiotherapy'/de OR 'resistance training'/de
#18	nondrug:ti,ab,kw OR 'non drug':ti,ab,kw OR nonpharmacolog*:ti,ab,kw OR 'non pharmacolog*':ti,ab,kw

#17	amitriptyline:ti,ab,kw OR amoxapine:ti,ab,kw OR clomipramine:ti,ab,kw OR desipramine:ti,ab,kw OR dothiepin:ti,ab,kw OR doxepin:ti,ab,kw OR imipramine:ti,ab,kw OR iprindole:ti,ab,kw OR lofepramine:ti,ab,kw OR nortriptyline:ti,ab,kw OR opipramol:ti,ab,kw OR protriptyline:ti,ab,kw OR trimipramine:ti,ab,kw OR gabapentin:ti,ab,kw OR sildenafil:ti,ab,kw OR vardenafil:ti,ab,kw
#16	'tricyclic antidepressant agent'/de OR 'amitriptyline'/de OR 'clomipramine'/de OR 'desipramine'/de OR 'dosulepin'/de OR 'doxepin'/de OR 'imipramine'/de OR 'iprindole'/de OR 'lofepramine'/de OR 'nortriptyline'/de OR 'opipramol'/de OR 'protriptyline'/de OR 'trimipramine'/de
#15	((antidepress*:ti,ab,kw OR 'anti depress*:ti,ab,kw OR '5 hydroxytryptophan':ti,ab,kw OR amisulpride:ti,ab,kw OR bupropion:ti,ab,kw OR citalopram:ti,ab,kw OR escitalopram:ti,ab,kw OR fluoxetine:ti,ab,kw OR fluvoxamine:ti,ab,kw OR maprotiline:ti,ab,kw OR mianserin:ti,ab,kw OR paroxetine:ti,ab,kw OR quipazine:ti,ab,kw OR ritanserin:ti,ab,kw OR sulpiride:ti,ab,kw OR trazodone:ti,ab,kw OR tryptophan:ti,ab,kw OR 'venlafaxine hydrochloride':ti,ab,kw OR viloxazine:ti,ab,kw OR ssri:ti,ab,kw OR ssris:ti,ab,kw OR 'selective serotonin reuptake':ti,ab,kw OR 'selective serotonin re-uptake':ti,ab,kw OR snri:ti,ab,kw OR snris:ti,ab,kw OR serotonin:ti,ab,kw) AND 'noradrenaline reuptake inhibitor':ti,ab,kw OR serotonin:ti,ab,kw) AND 'noradrenaline reuptake inhibitors':ti,ab,kw OR nri:ti,ab,kw OR nris:ti,ab,kw OR 'norepinephrine reuptake inhibitor':ti,ab,kw OR 'norepinephrine reuptake inhibitors':ti,ab,kw
#14	'antidepressant agent'/de OR 'serotonin uptake inhibitor'/de OR 'serotonin noradrenalin reuptake inhibitor'/de OR 'adrenergic receptor affecting agent'/de OR '5 hydroxytryptophan'/de OR 'amisulpride'/de OR 'amfebutamone'/de OR 'citalopram'/de OR 'fluoxetine'/de OR 'fluvoxamine'/de OR 'maprotiline'/de OR 'mianserin'/de OR 'paroxetine'/de OR 'quipazine'/de OR 'ritanserin'/de OR 'sulpiride'/de OR 'trazodone'/de OR 'tryptophan'/de OR 'venlafaxine'/de OR 'viloxazine'/de
#13	'cognitive behavior*:ti,ab,kw OR 'cognitive behaviour*:ti,ab,kw OR intervention:ti,ab,kw OR interventions:ti,ab,kw OR nondrug:ti,ab,kw OR 'non drug':ti,ab,kw OR nonpharmac*:ti,ab,kw OR 'non pharmac*:ti,ab,kw OR pharmac*:ti,ab,kw OR program*:ti,ab,kw OR psych*:ti,ab,kw OR psychosocial:ti,ab,kw OR 'psycho social':ti,ab,kw OR rehabilitation:ti,ab,kw OR therapy:ti,ab,kw OR therapies:ti,ab,kw OR treat*:ti,ab,kw
#12	'behavior therapy'/de OR 'cognitive behavioral therapy'/de OR 'mental health service'/de OR 'psychotherapy'/de OR 'psychosocial care'/de OR 'social support'/de OR 'motivational interviewing'/de OR 'patient participation'/de
#11	(checklist*:ti,ab,kw OR 'check list*:ti,ab,kw OR questionnaire:ti,ab,kw OR questionnaires:ti,ab,kw OR instrument:ti,ab,kw OR instruments:ti,ab,kw OR inventory:ti,ab,kw OR inventories:ti,ab,kw OR scale:ti,ab,kw OR scales:ti,ab,kw OR schedule:ti,ab,kw OR schedules:ti,ab,kw OR screen:ti,ab,kw OR screened:ti,ab,kw OR screening:ti,ab,kw OR 'beck depression':ti,ab,kw OR bdi:ti,ab,kw OR bdi2:ti,ab,kw OR 'geriatric depression scale':ti,ab,kw OR gds:ti,ab,kw OR 'hamilton rating scale for depression':ti,ab,kw OR 'hospital anxiety':ti,ab,kw) AND 'depression scale':ti,ab,kw OR hads:ti,ab,kw OR 'kidney disease quality of life':ti,ab,kw OR kdqol:ti,ab,kw OR 'medical outcomes study short form health survey 36':ti,ab,kw OR 'mos sf36':ti,ab,kw OR 'minnesota multiphasic personality inventory':ti,ab,kw OR mmpi:ti,ab,kw OR 'multiple affect adjective check list':ti,ab,kw OR maac:ti,ab,kw OR 'patient health questionnaire':ti,ab,kw OR phq2:ti,ab,kw OR 'phq 2':ti,ab,kw OR phq9:ti,ab,kw OR 'phq 9':ti,ab,kw OR 'prime-md':ti,ab,kw OR 'epidemiologic studies depression scale':ti,ab,kw OR ced:ti,ab,kw OR cesd:ti,ab,kw OR bref:ti,ab,kw OR dass21:ti,ab,kw OR idid:ti,ab,kw OR 'quick inventory of depressive symptomatology self-report':ti,ab,kw OR 'qids sr':ti,ab,kw OR 'rand 36-item health survey':ti,ab,kw OR 'rand 36':ti,ab,kw OR 'short form 36':ti,ab,kw OR 'sf 36':ti,ab,kw OR 'structured clinical interview':ti,ab,kw OR scid:ti,ab,kw OR 'self-rating depression scale':ti,ab,kw OR sds:ti,ab,kw OR 'short form health survey 36':ti,ab,kw OR sf36:ti,ab,kw
#10	'brief psychiatric rating scale'/de OR 'diagnostic procedure'/de OR 'neuropsychological test'/de OR 'patient health questionnaire'/de OR 'psychological rating scale'/de OR 'psychologic test'/de OR 'questionnaire'/de
#9	#5 AND #8
#8	#6 OR #7
#7	depressed:ti,ab,kw OR depressive:ti,ab,kw OR depression*:ti,ab,kw OR suicidal:ti,ab,kw OR suicide:ti,ab,kw OR suicides:ti,ab,kw
#6	'depression'/exp
#5	#1 OR #2 OR #3 OR #4
#4	dialysis:ti,ab,kw OR haemodiafiltration:ti,ab,kw OR hemodiafiltration:ti,ab,kw OR 'haemo diafiltration':ti,ab,kw OR 'hemo diafiltration':ti,ab,kw OR haemofiltration:ti,ab,kw OR hemofiltration:ti,ab,kw OR 'haemo filtration':ti,ab,kw OR 'hemo filtration':ti,ab,kw OR haemodialysis:ti,ab,kw OR hemodialysis:ti,ab,kw OR 'haemo dialysis':ti,ab,kw OR 'hemo dialysis':ti,ab,kw



#3	'hemodialysis'/exp OR 'hemofiltration'/exp OR 'hemodiafiltration'/exp OR 'peritoneal dialysis'/exp
#2	((((chronic OR endstage OR 'end stage' OR endstate OR 'end state' OR failure OR 'long term' OR maintenance) NEAR/2 (kidney OR renal)):ti,ab,kw) OR eskd:ti,ab,kw OR eskf:ti,ab,kw OR esrd:ti,ab,kw OR esrf:ti,ab,kw
#1	'chronic kidney failure'/exp

## EBM Reviews:

### Cochrane Central Register of Controlled Trials

### Cochrane Database of Systematic Reviews 2005 to June 24, 2020

### Database of Abstracts of Reviews of Effects 1st Quarter 2016

### Health Technology Assessment 4th Quarter 2016

Date searched: **June 24, 2020**

#	Searches
1	((((chronic or endstage or end-stage or endstate or end-state or failure or long-term or maintenance) adj2 (kidney or renal)) or ESKD or ESKF or ESRD or ESRF).ti,ab.
2	(dialysis or haemodiafiltration or hemodiafiltration or haemo-diafiltration or hemo-diafiltration or haemofiltration or hemofiltration or haemo-filtration or hemo-filtration or haemodialysis or hemodialysis or haemo-dialysis or hemo-dialysis).ti,ab.
3	or/1-2
4	(depressed or depressive or depression* or suicidal or suicide or suicides).ti,ab.
5	and/3-4
6	(checklist* or check-list* or questionnaire or questionnaires or instrument or instruments or inventory or inventories or scale or scales or schedule or schedules or screen or screened or screening or "Beck Depression" or BDI or BDI2 or "geriatric depression scale" or GDS or "Hamilton Rating Scale for Depression" or "Hospital Anxiety and Depression Scale" or HADS or "Kidney Disease Quality of Life" or KDQOL or "Medical Outcomes Study Short Form Health Survey 36" or MOS-SF36 or "Minnesota Multiphasic Personality Inventory" or MMPI or "Multiple Affect Adjective Check List" or MAACL or "Patient Health Questionnaire" or PHQ2 or PHQ-2 or PHQ9 or PHQ-9 or "PRIME-MD" or "Epidemiologic Studies Depression Scale" or CED or CESD or BREF or DASS21 or IDID or "Quick Inventory of Depressive Symptomatology Self-Report" or QIDS-SR or "RAND 36-Item Health Survey" or RAND-36 or "short form 36" or SF-36 or "Structured Clinical Interview" or SCID or "self-rating depression scale" or SDS or "Short Form Health Survey 36" or SF36).ti,ab.
7	(cognitive-behavior* or cognitive-behaviour* or intervention or interventions or nondrug or non-drug or nonpharmac* or non-pharmac* or pharmac* or program* or psych* or psychosocial or psycho-social or rehabilitation or therapy or therapies or treat*).ti,ab.
8	(antidepress* or anti-depress* or 5-hydroxytryptophan or Amisulpride or Bupropion or Citalopram or Escitalopram or Fluoxetine or Fluvoxamine or Maprotiline or Mianserin or Paroxetine or Quipazine or Ritanserin or Sulpiride or Trazodone or Tryptophan or Venlafaxine Hydrochloride or Viloxazine or SSRI or SSRIs or "selective serotonin reuptake" or "selective serotonin re-uptake" or SNRI or SNRIs or "Serotonin and Noradrenaline Reuptake Inhibitor" or "Serotonin and Noradrenaline Reuptake Inhibitors" or NRI or NRIs or "norepinephrine reuptake inhibitor" or "norepinephrine reuptake inhibitors").ti,ab.
9	(Amitriptyline or Amoxapine or Clomipramine or Desipramine or Dothiepin or Doxepin or Imipramine or Iprindole or Lofepramine or Nortriptyline or Opipramol or Protriptyline or Trimipramine or Gabapentin or Sildenafil or Vardenafil).ti,ab.
10	(nondrug or non-drug or nonpharmacolog* or non-pharmacolog*).ti,ab.
11	((((aerobic or resistance) adj2 (exercis* or program* or therap* or train*)) or (exercise adj2 (program* or therap* or train*)) or cross-training).ti,ab.
12	music therapy.ti,ab.
13	or/6-12
14	and/5,13

## ClinicalTrials.gov

Date searched: **June 24, 2020**

( depressed OR depressive OR depression\* OR suicidal OR suicide OR suicides ) AND INFLECT EXACT ( "Active, not recruiting" OR "Completed" OR "Suspended" OR "Terminated" OR "Withdrawn" OR "Unknown status" ) [OVERALL-STATUS] AND ( ( chronic OR endstage OR end-stage OR failure ) AND ( kidney OR renal ) OR ESKD OR ESKF OR ESRD OR ESRF OR dialysis OR haemodiafiltration OR hemodiafiltration OR haemo-diafiltration OR hemo-diafiltration OR haemofiltration OR hemofiltration OR haemo-fi ) [DISEASE]

## WHO ICTRP

Date searched: **June 24, 2020**

((((chronic OR endstage OR end-stage OR failure) AND (kidney OR renal)) OR ESKD OR ESKF OR ESRD OR ESRF) AND (depress\* OR suicid\*))

## VA HSR&D

<https://www.hsr.d.research.va.gov/research/>

Date searched: **June 24, 2020**

Separately searched terms: kidney and renal. Reviewed result lists

**Table S1. PICOTS by Key Question**

Key Question:	KQ1: What are the performance characteristics of screening tools for depression in patients with kidney failure?	KQ2: What is the impact of screening for depression in patients with kidney failure on intermediate and/or patient outcomes?	KQ4: In patients with ESRD, what are the potential harms of screening?	KQ5: Do the benefits or harms of screening differ by: a. patient characteristics or other social determinants of health? b. setting? c. screening characteristics/ process? d. other (e.g., patient engagement/ receptivity to treatment, social support)? e. Timing and type of follow up?
<b>Population</b>	Adults with kidney failure			
<b>Intervention</b>	Depression screening			
<b>Comparators</b>	Clinical evaluation, Other screening tools. Exclude DSM-III and earlier	No screening, other screening tool.		
<b>Outcomes</b>	<u>Diagnostic test performance:</u> sensitivity, specificity, positive predictive value, and negative predictive value	<u>Therapeutic impact:</u> timing, setting, or type of treatment. <u>Intermediate and Patient outcomes:</u> depressive symptoms, mortality, suicide attempts or completion, hospitalization, ED/urgent care utilization, patient satisfaction, adherence to dialysis, medication, or treatment, pain	Adverse effects or unintended consequences	<u>Intermediate and Patient outcomes:</u> depressive symptoms, mortality, suicide attempts or completion, hospitalization, ED/urgent care utilization, patient satisfaction, adherence to dialysis, medication, or treatment, pain medication reduction, BP/metabolic control, quality of life, other outcomes (e.g., employment)

		medication reduction, BP/metabolic control, quality of life, other outcomes (e.g., employment)		
<b>Timing</b>	Any			
<b>Settings</b>	All settings in U.S. or international (VHA, hospital community, community mental health, ED, urgent care, other community)			
<b>Study design</b>	Systematic reviews, RCTs, NRCTs, Observational studies			

Note. Subpopulations may include: Patient demographic characteristics or social determinants of health; ESRD subgroup (w/o treatment; treated by kidney transplant; treated by HD (home or clinic); treated by PD (home or clinic); clinical severity (ESRD or depression); setting (*eg* VHA, community hospitals, community mental health, ED, urgent care visits for mental health, home vs clinic-based dialysis); other.

Abbreviations: CES-D = Center for Epidemiologic Studies Depression Scale; BDI = Beck Depression Inventory; BP = blood pressure; ED = emergency department; ESRD = End-stage Renal Disease; HADS = Hospital Anxiety and Depression Scale; HAM-D = Hamilton Depression Rating Scale; NRCT = Non-randomized controlled trial; PHQ-9 = Patient Health Questionnaire-9; RCT = Randomized controlled trial; VHA = Veterans Health Administration

**Table S2. QUADAS-2 Risk of Bias Assessment**

	Could the selection of patients have introduced bias?				Could the conduct or interpretation of the index test have introduced bias?					Could the conduct or interpretation of the reference standard have introduced bias?					Could the patient flow have introduced bias?				Applicability			
	1	2	3	ROB	4	5	6	7	ROB	8	9	10	11	ROB	12	13	14	ROB	15	16	17	ROB
Alsuwaida, 2006(1)	U	Y	U	Unclear	Y	U	U	Y	Unclear	Y	Y	U	Y	Unclear	Y	NA	U	Unclear	N	N	N	Low
Balogun, 2011(2)	U	Y	Y	Unclear	Y	U	U	Y	Unclear	Y	U	U	U	Unclear	Y	Y	N	High	N	N	N	Low
Bautovich, 2018(3)	U	Y	Y	Unclear	Y	U	NA	Y	Unclear	Y	Y	N	Y	High	Y	NA	Y	Low	N	N	N	Low
Chilcot, 2008(4)	U	Y	Y	Unclear	U	U	NA	Y	Unclear	U	Y	U	U	Unclear	Y	NA	Y	Low	N	N	N	Low
Collister, 2019(5)	Y	Y	Y	Low	U	U	U	Y	Unclear	U	U	U	U	Unclear	Y	NA	Y	Low	N	N	N	Low
Gencoz, 2007(6)	Y	Y	U	Unclear	Y	Y	Y	Y	Low	Y	U	Y	U	Unclear	Y	Y	Y	Low	N	N	N	Low
Giordano, 2007(7)	Y	U	Y	Unclear	U	Y	Y	Y	Unclear	Y	Y	U	Y	Unclear	Y	NA	N	High	N	N	N	Low
Grant, 2008(8)	U	Y	Y	Unclear	U	U	NA	Y	Unclear	U	Y	U	Y	Unclear	Y	NA	Y	Low	N	N	N	Low
Hedayati, 2006(9)	U	Y	Y	Unclear	Y	U	NA	Y	Unclear	U	Y	U	Y	Unclear	Y	NA	Y	Low	N	N	N	Low
Loosman, 2010(10)	U	Y	Y	Unclear	Y	U	NA	Y	Unclear	Y	Y	Y	Y	Low	Y	NA	Y	Low	N	N	N	Low
Neitzer, 2012(11)	U	Y	Y	Unclear	U	U	NA	Y	Unclear	U	U	NA	Y	Unclear	Y	NA	Y	Low	N	N	N	Low
Preljevic, 2012(12)	U	Y	Y	Unclear	Y	U	NA	Y	Unclear	Y	U	Y	Y	Unclear	Y	NA	N	Low	N	N	N	Low
Troidle, 2003(13)	Y	Y	Y	Low	N	U	U	Y	High	U	U	U	U	Unclear	Y	NA	Y	Low	N	N	N	Low
Van den Beukel, 2012(14)	U	Y	Y	Unclear	U	U	U	Y	Unclear	U	U	U	Y	Unclear	Y	NA	Y	Low	N	N	N	Low
Wang, 2019(15)	Y	Y	Y	Low	Y	U	U	Y	Unclear	Y	U	U	Y	Unclear	Y	NA	Y	Low	N	N	N	Low
Watnick, 2005(16)	U	Y	Y	Unclear	Y	U	NA	Y	Unclear	Y	Y	U	Y	Unclear	Y	NA	Y	Low	N	N	N	Low

Abbreviations: N = no; NA = not applicable; ROB = risk of bias; U = unclear; Y = yes

Questions (QUADAS-2(17)):

1. Consecutive or random sample of patients enrolled?
2. Was a case-control design avoided?
3. Did the study avoid inappropriate exclusions?
4. Was the index test interpreted without knowledge of the reference standard results?
5. Was staff trained in the use of the index test?

6. Was the fidelity of the index test monitored and/or reported?
7. Is the reference standard likely to correctly classify the target condition?
8. Was the reference standard interpreted without knowledge of the index test results?
9. Was staff trained in the assessment of the reference standard?
10. Was the fidelity of the reference test monitored and/or reported?
11. Was there an appropriate interval between the index test and reference standard?
12. Did all patients receive the same reference standard?
13. If a partial selection of patients received the reference standard, was it adjusted?
14. Were all patients included in the analysis?
15. Are there concerns that the study population differs from the review question?
16. Are there concerns that the index test, its conduct, or its interpretation differ from the review question?
17. Are there concerns that the reference standard, its conduct, or its interpretation differ from the review question?

**Table S3. Characteristics of studies examining the diagnostic accuracy of depression screening tools in patients with kidney failure**

<b>Author, Year N enrolled Country/US region Years of enrollment</b>	<b>Study setting Sample characteristics and demographics</b>	<b>Study Inclusion/Exclusion Criteria</b>	<b>Index Test(s): Test Administration</b>	<b>Reference Standard: Test Administration</b>
Alsuwaida, 2006(1) N = 26 Saudi Arabia	Single Site: hospital-based outpatient HD unit 42% Female Age: 48.1(15.1) Education: NR HD: 100% Dialysis duration: NR History of depression: NR	Inclusion: 18+ years of age, kidney failure and on maintenance HD for 3+ months Exclusion: Inability to participate in psychiatric interview, acute kidney failure, and delirium. Diagnosed with psychiatric disorders other than MDD	<u>SRQ (Arabic Version):</u> Self-report. Timing: within a week of clinical interview	<u>Clinical Interview:</u> All participants interviewed by the same psychiatrist (blinded to index test). Timing: up to a week before the index test.
Balogun, 2011(2) N = 96 US	Multisite: dialysis units Of 89 participants: 56% Female Age: 73.5(6.2) White: 56.2% Black: 43.8% Education: NR HD: NR Dialysis duration: NR History of depression: NR	Inclusion: 65+ with kidney failure treated with maintenance hemodialysis and able to give their informed consent Exclusion: acute or other chronic illness [ <i>i.e.</i> , metabolic (organic) brain syndrome, known malignancy, dementia], currently using antidepressants, and active alcohol or recreational drug abuse, did not speak English	<u>BDI-II, GDS-15:</u> NR	<u>Clinical Interview:</u> Geriatric Psychiatrist
Bautovich, 2018(3) N = 45 Sydney, Australia	Single site: outpatient dialysis unit 42% Female Age: primarily 65+ Education: NR HD: 100% Days on dialysis: M= 1241(1098) History of depression: NR	Included: 18+ years of age, receiving HD, adequate English language skills Excluded: evidence of psychosis, drug or alcohol dependence, or cognitive dysfunction	<u>BDI-II, CDI:</u> Self-report. Timing: before clinical interview.	<u>Clinical Interview:</u> Interviewed by a senior psychiatry registrar or psychiatrist, both of whom were experienced in diagnosing depression amongst those with chronic medical illness;

Author, Year N enrolled Country/US region Years of enrollment	Study setting Sample characteristics and demographics	Study Inclusion/Exclusion Criteria	Index Test(s): Test Administration	Reference Standard: Test Administration
				Timing: completed immediately after index tests
Chilcot, 2008(4) N = 40 UK	Multisite: outpatient renal service 40% Female Age: 53.2(14.2) White: 87.5% Black Caribbean: 10% Asian: 2.5% Education: NRHD: 100% (high-flux or on-line) 3x/week Months on dialysis M=51.2 History of depression: NR	Included: Adult kidney failure receiving HD for >3 months. Excluded: Psychiatric illnesses other than MDD, <23 MMSE	<u>BDI-II, CDI:</u> Self-report. Completed on and off dialysis. On-dialysis commenced 30 minutes after the start of a stable session. Off-dialysis conducted at the same as the MINI, M=10.7(4.2) days before/after.	<u>MINI:</u> Administration by a research psychologist who was trained by a consultant psychiatrist. Timing: 10.7(4.2) days before/after the on-dialysis BDI-II, and on the same day as the off-dialysis BDI-II.
<sup>1</sup> Collister, 2019(5) N = 50 Canada	Multisite: outpatient HD units 48% Female Age: 64(12.4) Education: NR HD: 100% 3+x/week: 96% Hours of HD M= 3.6(0.4) Dialysis duration: NR History of depression: NR Antidepressants: 16%	Included: 18+ years of age, receiving in-center hemodialysis ≥2x weekly for at least the last 90 days Excluded: unable to complete the study instruments due to a cognitive impairment or an English language barrier	<u>Single question from the ESAS:</u> Self-report scale (0-10) re: feeling blue or sad. Timing: taken during dialysis during the same session as the reference test.	<u>HADS:</u> Self-report. Timing: taken during dialysis during the same session as the reference test.
<sup>1</sup> Gencoz, 2007(6) N = 45 Turkey	Single site: hospital-based outpatient HD unit 42.2% Female Age: 41.64(11.7) ≤ Middle school: 37.9%	Included: medically stable with no hospital admission for any reason within the last 3 months, and maintained on dialysis for at least 12 months	<u>Ham-D:</u> Administered at baseline and the following month by a clinical psychologist that was blind to the	<u>SCID-I (Turkish Translation):</u> Administered at baseline and the following month by a clinical psychologist



Author, Year N enrolled Country/US region Years of enrollment	Study setting Sample characteristics and demographics	Study Inclusion/Exclusion Criteria	Index Test(s): Test Administration	Reference Standard: Test Administration
	HD: 100% Months on HD M=72.24(48.48) History of depression: NR	Excluded: presence of cognitive impairment indicated by MMSE score lower than 24, presence of a history of a psychiatric diagnosis or treatment in the last 6 months, and presence of some practical difficulties like probability of moving to another city, blindness or low educational level, which may decrease the patients' ability to comprehend and/or follow the study protocol. Patients who did not complete all baseline assessments were also excluded from the study.	reference standard. Timing re: reference standard: NR	that was blind to the reference standard. Timing re: index test: NR
<sup>1</sup> Giordano, 2007(7) N = 31 Italy	Single site: hospital-based HD unit 35.5% Female Age: 70.3(1) Race: NR Education: NR HD: 100% Dialysis duration: NR History of depression: NR	Inclusion: 3+ HD/wk, 65+ years old, maintaining functional independence or loss of it in only 1 of the 6 basic ADL, no evidence of significant cognitive impairment per MMSE >24, no evidence of severe diseases that might highly influence mood state (e.g., cancer, symptomatic cerebrovascular disease with residual deficit, schizophrenia and other psychoses), and disease severity as evaluated by the CIRS for overall illness severity for which >3 is moderate Exclusion: Taking antidepressants	<u>GDS-15</u> : Self-report. Administered by a trained interviewer. Timing: same session as reference standard.	<u>BDI-II</u> : Self-report. Administered by a trained interviewer. Timing: same session as index test.

<b>Author, Year N enrolled Country/US region Years of enrollment</b>	<b>Study setting Sample characteristics and demographics</b>	<b>Study Inclusion/Exclusion Criteria</b>	<b>Index Test(s): Test Administration</b>	<b>Reference Standard: Test Administration</b>
<sup>2</sup> Grant, 2008(8) N = 57 UK	Single site: outpatient HD unit 29.8% Female Age: 62.5(15.8) Non-White 7% Education: NR HD: 100% Dialysis duration: NR History of depression: NR	Included: 18 and 90 years of age, kidney failure for 3+ months, receiving HD 3x/week. Excluded: Current psychiatric care, on medication for a psychiatric illness or had seen a psychiatrist for follow-up within the last 2 years, severe co- morbid illness requiring hospitalization.	<u>BDI-II:</u> Self-report. Distributed by a healthcare assistant during a HD session.	<u>Clinical Interview (based on ICD-10 diagnosis):</u> Interviewed by a trained psychologist. Included a full psychiatric history and MMSE. Timing: within 1 week of index test
<sup>1</sup> Hedayati, 2006(18) N = 98 US Durham, NC March 2003- April 2004	Multi-site: outpatient dialysis units (VA, 2 non-VA) 44.9% Female Age: 57.2(13.8) Veterans: 26.5% AA/Black: 80.6% White: 14.3% Other: 5.1% ≤High school: ≈ 44.5% HD: 100% Years on dialysis: M=4.1(3.8) History of depression: NR	Included: English-speaking with health- care power of attorney and could sign consent. Excluded: NR	<u>BDI-II, CDI, CESD, Feinstein Scale:</u> RA administered BDI- II/CESD/Feinstein Scale at enrollment.	<u>SCID-I:</u> Administered by a nephrologist. Timing: within 1 week of index tests
<sup>1</sup> Loosman, 2010(10) N = 62 Amsterdam Feb-June 2008	Single site: hospital-based HD and outpatient PD 46.8% Female Age: 63.5(14.9) 64.5% Dutch ethnicity Education: NR HD: 82%; PD 18% Months on dialysis: 46(65) Previous depression: 9.7% Antidepressants: 3.2%	Included: Patients with kidney failure treated with HD or PD Excluded: Patients who were unable to read or understand Dutch	<u>BDI-II, HADS:</u> Self-report. Completed while receiving treatment.	<u>MINI:</u> Performed by a medical resident who was extensively trained on the MINI by a psychiatrist. For 1:7 patients, MINI interviews were performed by both the medical resident and the psychiatrist (100% Inter-

Author, Year N enrolled Country/US region Years of enrollment	Study setting Sample characteristics and demographics	Study Inclusion/Exclusion Criteria	Index Test(s): Test Administration	Reference Standard: Test Administration
Neitzer, 2012(11) N = 134 US CA, TX 2009	Multisite: outpatient HD units 48% Female Age: 59.1(14.7) AA/Black: 22% Asian: 13% White: 60% Other: 4% Education: NR HD: 100% Months on dialysis: Median = 27.5 (2.9-252.2) History of depression: NR	Included: English or Spanish speaking, 18+ years old, due in April to June 2009 for their KDQOL-SF36 assessment. Excluded: Questionnaires with 50% or more of the questions left blank were considered incomplete and excluded.	<u>BDI-FS:</u> Self-report. Completed during HD treatment.	rater reliability). Timing: NR  <u>BDI-II:</u> Completed during HD session. Order of completion was not specified.
<sup>1</sup> Preljevic, 2012(12) N = 109 Norway	Multisite: hospital-based HD and PD centers 30.3% Female Age 57.8(15.7) Race: NR 69.4% HS or less HD: 76.6%; PD: 23.3% Months on dialysis: M=8.5 (3.75–22) History of depression: NR	Included: 18+ years receiving either HD or PD for more than 2 months, were in a stable clinical condition and had adequate Norwegian language skills. Excluded: Cognitive dysfunction, psychosis or drug/alcohol abuse; hospitalization during the investigation period; however, they could be enrolled 4 weeks or more after discharge from hospital if they were in a stable clinical condition.	<u>BDI-II, CDI, HADS-D:</u> Self-report. Completed in a standardized sequence during the dialysis treatment for HD patients and during the routine outpatient control for PD patients.	<u>SCID-I:</u> Administered by an experienced psychiatrist who was blinded to each participant's medical history and scores on all self-report questionnaires. Assessments were conducted during dialysis sessions to standardize the assessment procedure and the time point relative to dialysis treatment. Interviews were

Author, Year N enrolled Country/US region Years of enrollment	Study setting Sample characteristics and demographics	Study Inclusion/Exclusion Criteria	Index Test(s): Test Administration	Reference Standard: Test Administration
<sup>1</sup> Troidle, 2003(13) N = 97 US June 2000 – January 2002	Multisite: CPD and HD units CP: 46% Female; HD: 40% Female Age: CPD 55.4(11.3); HD 56(8.6) White: CPD 75%; HD 87% CPD: 83%; HD 17% Education: NR Dialysis duration: NR History of depression: NR	NR	<u>2 items from the KDQOL SF-36:</u> Self-report. Likert 1-6. Scored by a social worker. Timing: consecutive	audiotaped and 25 randomly selected tapes were scored independently by another psychiatrist to establish inter-rater reliability. The interrater reliability for depressive disorder was excellent ( $\kappa=1$ ). Timing: NR
<sup>1</sup> Van den Beukel,(14) 2012 N = 133 Netherlands	Multisite: outpatient hospital- based dialysis units 39% Female Age: 62(16) Native Dutch: 66% Education: NR HD: 72% Dialysis duration: NR Previous Depression: 9% Antidepressant: 6% Months on dialysis: M=54(65)	Inclusion: 18+ years of age, kidney failure for at least 30 days, able to read the Dutch language and had no significant visual, physical, or cognitive impairment that would prevent completion of the questionnaires Exclusion: NR	<u>MHI5 of the SF-36:</u> Self-report. Completed during dialysis. Timing: NR	<u>BDI-II/CDI (Dutch Translation):</u> Self-report. Completed during dialysis. Timing: NR

Author, Year N enrolled Country/US region Years of enrollment	Study setting Sample characteristics and demographics	Study Inclusion/Exclusion Criteria	Index Test(s): Test Administration	Reference Standard: Test Administration
	History of depression: NR			
Watnick, 2005(16) N = 62 US Portland, OR July 2003- May 2004	Multisite: public and private outpatient HD and PD units (including VA) Female: 32% Age: 63(15) AA/Black: 15% Hispanic: 5% Asian: 5% White: 76% Education: NR HD: 95%, PD: 5% Dialysis duration: NR History of depression: NR	Inclusion: 18+ years old and had started dialysis therapy more than 90 days before enrollment. Exclusion: Did not speak English, MMSE $\leq$ 17, medical record documentation of a psychiatric diagnosis other than depression, were deemed unable to participate by the dialysis staff, or were scheduled for kidney transplant within the next month.	<u>BDI-II, PHQ-9:</u> Self-report.	<u>SCID-I:</u> Interviewed by a mental health professional (completed psychology internship), blinded to BDI- II/PHQ-9 results. Timing: within 2 weeks of index tests.
<sup>1</sup> Wang, 2019(15) N = 319 China	Multisite: hospital-based HD units 31.4% Female depressed; 43.78% Female non- depressed Age: 49.4 (6.04) depressed; 50.92(6.46) non-depressed Race: NR HS or less: 51.44% depressed; 57.78% non-depressed HD:100% Dialysis duration: NR History of depression: NR	Inclusion: 18+ years of age; history of maintenance HD >3 months; ability to understand written Chinese, complete the interview and the questionnaire, and provide informed consent Excluded: documented cognitive impairment, had another primary diagnosis (e.g. chronic heart failure, cancer, hyperthyroidism), or had been previously diagnosed with depression and other psychiatric disorders	<u>BDI-II, DI-MHD:</u> Self-report. Timing: 2 weeks after clinical interview	<u>SCID-I:</u> Administered by a psychologist and a nephrologist. Timing: 2 weeks before index tests.

<sup>1</sup> Grouped milder forms of depression with major depressive disorder. <sup>2</sup> Included cutoff values for both major depressive disorder as well as for milder forms of depression and subclinical symptoms.

Abbreviations: BDI-II = Beck Depression Inventory – II; BDI-FS = Beck Depression Inventory - Fast Screen; CA = California; CDI = Cognitive Depression Index; CES-D = Center for Epidemiologic Studies – Depression Scale; CVD = cardiovascular disease; DI-MHD = Depression Inventory – Maintenance Hemodialysis; ESAS = Edmonton Symptom Assessment System; GDS-15= Geriatric Depression Scale-15 ; HADS-D = Hospital Anxiety and Depression Scale - Depressive Subscale; Ham-D= Hamilton Depression Rating Scale; HD = hemodialysis; HS = high school; ICD-10 = 10th revision of the International Statistical Classification of Diseases and Related Health Problems; KDQOL SF-36 = Kidney Disease Quality of Life Short Form - 36; MDD = major depressive disorder; MHI5 = Mental Health Inventory 5; NR = not reported; MINI = Mini International Neuropsychiatric Interview; MMSE = Mini-Mental Status Examination; NR = Not reported; NC = North Carolina; OR = Oregon; PD = peritoneal dialysis; PHQ-9= Patient Health Questionnaire 9; SCID-I= Structured Clinical Interview for DSM-IV; SF-36 = Kidney Disease Quality of Life Short Form - 36; SRQ = Self-Reporting Questionnaire; TX = Texas; UK = United Kingdom; US = United States; VA = Veterans Affairs

**Table S4. Findings of studies examining the diagnostic accuracy of depression screening tools in patients with kidney failure compared to a gold standard diagnostic interview**

Author, Year N enrolled	Cut-off	Sens (%)	Spec (%)	PPV (%)	NPV (%)	AUC	Summary of Findings
<b>Beck Depression Inventory-II (BDI-II)</b>							
Balogun, 2011(2) N = 96	≥10	68	77	57	85	0.73	Compared to diagnostic interview, the BDI-II cutoff with the best diagnostic accuracy was ≥10.
Bautovich, 2018(3) N = 45	≥18	100	90	60	100	0.99	Compared to diagnostic interview, the BDI-II is an acceptable screening tool, with a cutoff of ≥18.
Chilcot, 2008(4) N = 40	≥16	88.9	87.1	88.8	87	0.961	Consistent with previous research, (off dialysis) BDI-II with a cutoff of ≥16 has good diagnostic accuracy.
<sup>2</sup> Grant, 2008(8) N = 57	≥10	100	50	21.9	100	0.93	Using the general population cut-off score (≥10), the BDI-II significantly over-diagnosed depression in this HD population. A cutoff of ≥15 is more reliable.
	≥15	100	78	NR	NR	0.93	
	≥20	71.4	92	NR	NR	0.93	
<sup>1</sup> Hedayati, 2006(9) N = 98	≥14	62	81	53	85	0.77	When used for screening, the threshold for depression should be higher for patients with versus without kidney failure (i.e., ≥14).
<sup>1</sup> Loosman, 2010(10) N = 62	≥13	75	90.2	75	90.2	0.90	At a cutoff of ≥13, the BDI-II is an effective screening tool for depression in depression in patients with kidney failure.
<sup>1</sup> Preljevic, 2012(12) N = 109	≥12	91	63	39	96	0.92	The BDI-II demonstrated acceptable performance as a screening tool for depression. At a threshold of ≥16 (general population) the BDI-II performed better than the HADS and the CDI; however, a cutoff of ≥17 is more reliable for this population.
	≥13	91	68	43	97	0.92	
	≥14	86	71	44	95	0.92	
	≥15	82	75	46	94	0.92	
	≥16	82	87	62	95	0.92	
	≥17	82	89	67	95	0.92	
	≥18	77	92	71	94	0.92	
<sup>1</sup> Wang, 2019(15) N = 319	≥15	87	49	34	93	0.84	A cutoff of ≥19 indicated depression in this population.
	≥16	87	58	39	94	0.84	
	≥17	87	65	43	94	0.84	
	≥18	87	71	47	95	0.84	
	≥19	83	86	63	94	0.84	
	≥20	74	94	77	92	0.84	
Watnick, 2005(16) N = 62	≥16	91	86	59	98	0.937	The BDI-II ≥16 is a valid measure for depressive disorders in the dialysis population.
<b>Cognitive Depression Index (CDI)</b>							
Bautovich, 2018(3)	≥11	100	92	67	10	0.98	Compared to diagnostic interview, the CDI ≥11 is an acceptable screening tool.

Author, Year N enrolled	Cut-off	Sens (%)	Spec (%)	PPV (%)	NPV (%)	AUC	Summary of Findings
N = 45							
Chilcot, 2008(4) N = 40	≥10	77.8	80.6	77.7	80.6	0.94	The CDI ≥10 has reduced sensitivity and specificity as compared to the BDI-II.
<sup>1</sup> Hedayati, 2006(9) N = 98	≥8	50	83	52	82	0.76	When used for screening, the threshold for depression should be higher for patients with versus without kidney failure. The BDI-II or the CESD have better sensitivity and better agreement (kappa) than the CDI (cutoff ≥8).
<sup>1</sup> Preljevic, 2012(12) N = 109	≥9	82	79	50	94	0.89	The CDI (cutoff ≥11) demonstrated acceptable performance as a screening tool for depression. The BDI-II performed better than the CDI.
	≥10	82	86	60	95	0.89	
	≥11	82	93	75	95	0.89	
	≥12	77	95	81	94	0.89	
	≥13	50	98	85	88	0.89	
	≥14	41	98	82	86	0.89	
<b>Center for Epidemiologic Studies – Depression Scale (CES-D)</b>							
<sup>1</sup> Hedayati, 2006(9) N = 98	≥18	69	83	60	88	0.86	When used for screening, the CESD threshold for depression should be higher (≥18) for patients with versus without kidney failure.
<b>Depression Inventory – Maintenance Hemodialysis (DI-MHD) – Chinese Language</b>							
<sup>1</sup> Wang, 2019(15) N = 319	≥23	97	55	84	89	0.94	At a threshold of ≥25, the DI-MHD performed better than the BDI-II.
	≥24	97	72	90	91	0.94	
	≥25	97	86	95	93	0.94	
	≥26	93	90	96	84	0.94	
	≥27	85	90	95	70	0.94	
<b>Geriatric Depression Scale-15 (GDS-15)</b>							
Balogun, 2011(2) N = 96	≥5	63	82	60	83	0.81	The GDS-15 ≥5 is a valid tool compared to the gold standard.
<b>Hamilton Depression Rating Scale (Ham-D)</b>							
<sup>1</sup> Gencoz, 2007(6) N = 45	≥10	100	80	59	100	85	The HDRS ≥10 is a reliable and valid instrument that can be used among patients with kidney failure undergoing HD
<b>Hospital Anxiety and Depression Scale - Depressive Subscale (HADS-D)</b>							
<sup>1</sup> Loosman, 2010(10) N = 62	≥6	90.5	75.6	85.7	75.6	0.89	The HADS-D ≥6 is an effective screening tool for depression in depression in patients with kidney failure.
<sup>1</sup> Preljevic, 2012(12) N = 109	≥4	100	48	33	100	0.91	At a HADS-D threshold of ≥8 the BDI-II performed better.
	≥5	95	60	38	98	0.91	
	≥6	95	73	48	98	0.91	
	≥7	86	84	58	96	0.91	



Author, Year N enrolled	Cut-off	Sens (%)	Spec (%)	PPV (%)	NPV (%)	AUC	Summary of Findings
	≥8	73	87	59	93	0.91	
	≥9	59	92	65	90	0.91	
	≥10	59	94	72	90	0.91	
	≥11	50	96	79	88	0.91	
<b>Patient Health Questionnaire 9 (PHQ-9)</b>							
Watnick, 2005(16) N = 62	≥10	92	92	71	98	0.94	The PHQ-9 ≥10 is a valid measure for depressive disorders in the dialysis population.
<b>Self-Reporting Questionnaire (SRQ)</b>							
Alsuwaida, 2006(1) N = 26	≥8	100	50	26	NR	0.96	The SRQ has high sensitivity the PPV is poor due to the somatic symptoms in non-depressed patients with kidney failure. A cutoff of ≥13 results in an acceptable specificity level without compromising sensitivity.
	≥9	100	68	36	NR	0.96	
	≥11	100	72	40	NR	0.96	
	≥13	100	82	50	NR	0.96	
	≥14	75	91	60	NR	0.96	
	≥16	75	95.5	75	NR	0.96	
≥18	75	100	100	NR	0.96		

<sup>1</sup> Screened for depressive symptoms or milder forms of depression in addition to Major Depressive Disorder.

<sup>2</sup> Included cutoff values for both Major Depressive Disorder as well as for milder forms of depression and subclinical symptoms.

Abbreviations: AUC = Area under (receiver operating characteristic [ROC]) curve; BDI-II = Beck Depression Inventory – II; CDI = Cognitive Depression Index; CES-D = Center for Epidemiologic Studies – Depression Scale; GDS-15 = Geriatric Depression Scale-15; HADS-D = Hospital Anxiety and Depression Scale - Depressive Subscale; Ham-D = Hamilton Depression Rating Scale; NPV = Negative predictive value; NR = Not reported; PHQ-9 = Patient Health Questionnaire 9; PPV = Positive predictive value; Sens = sensitivity; Spec = specificity; SRQ = Self-Reporting Questionnaire

**Table S5. Studies Comparing a Depression Tool to Another Validated Depression Tool**

Author, Year N enrolled	Cutoff	Sens (%)	Spec (%)	PPV (%)	NPV (%)	AUC	Summary of Findings
Beck Depression Inventory – Fast Screen (BDI-FS)							
Neitzer, 2012(11) N = 134	≥4	97.2	91.8	81.4	98.9	0.98	Reference standard: BDI-II ≥16
Edmonton Symptom Assessment System (ESAS) – single question							
<sup>1</sup> Collister, 2019(5) N = 50	≥2	81	74	NR	NR	0.81	Reference standard: HADS ≥6.
Geriatric Depression Scale-15 (GDS-15)							
<sup>1</sup> Giordano, 2007(7) N = 31	≥6	89	92	NR	NR	0.95	Reference standard: BDI-II ≥14.
Kidney Disease Quality of Life Short Form - 36 (KDQOL SF-36) “Have you felt downhearted and blue?”							
<sup>1</sup> Troidle, 2003(13) N = 97	≤3	82	69	NR	NR	NR	Reference standard: BDI-II ≥11.
Kidney Disease Quality of Life Short Form - 36 (KDQOL SF-36) “Have you felt so down in the dumps so that nothing could cheer you?”							
<sup>1</sup> Troidle, 2003(13) N = 97	≤3	65	67	NR	NR	NR	Reference standard: BDI-II ≥11.
Mental Health Inventory 5 (MHI5)							
<sup>1</sup> Van den Beukel, 2012(14) N = 133	≥66	67	78	NR	NR	0.82	Reference standard: BDI-II ≥16 (≥66+) and CDI ≥10 (≥74+)
	≥70	77	72	44	91	0.82	
	≥74	83 81 CDI	65 65 CDI	NR	NR	0.82 0.81 CDI	
	≥78	90	54	NR	NR	0.82	
	≥82	93	45	NR	NR	0.82	

<sup>1</sup>Screened for depressive symptoms or milder forms of depression in addition to Major Depressive Disorder.

Abbreviations: AUC = Area under (receiver operating characteristic[ROC]) curve; BDI-II = Beck Depression Inventory – II; BDI -FS = Beck Depression Inventory - Fast Screen; ESAS = Edmonton Symptom Assessment System; GDS-15 = Geriatric Depression Scale-15; HADS-D = Hospital Anxiety and Depression Scale - Depressive Subscale; KDQOL-36 = Kidney Disease Quality of Life Short Form - 36; MHI5 = Mental Health Inventory 5; NPV = Negative predictive value; NR = Not reported; PHQ-9 = Patient Health Questionnaire 9; PPV = Positive predictive value; Sens = sensitivity; Spec = specificity; SRQ = Self-Reporting Questionnaire

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