Depression and quality of life in patients with chronic wounds: ways to measure their influence and their effect on daily life

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Abstract: Depression is a very common comorbidity in patients with chronic wounds. At least 30% of those patients suffer from depressive symptoms or anxiety. To measure impairment on daily life, different quality of life (QoL) questionnaires can be used as well as specialized questionnaires to evaluate different levels of depression. In this review, we present some of the current measurement tools and effects of depression on QoL. Depression is often associated with wound duration, odor of wounds, and furthermore with pain and immobility. On the other hand, depression might reduce wound healing itself via psychoimmunological effects. QoL, in general, is reduced in patients with chronic wounds, but especially in the case of pain, immobility and older age, and sometimes independent to wound status (healed or active). Treating medical and health care personnel should establish depression and QoL questionnaires in daily practice and should consider the psychological health of their patients as part of their treatment.

Keywords: quality of life questionnaire, depression assessments, pain, chronic wounds, anxiety, depression

Introduction
Depression is not only defined as an unhappy feeling, but in fact, is a widespread mental disorder characterized by ongoing sadness and depressed mood, anhedonia, loss of energy, appetite and/or interest in daily activities, guilt, loss of self-esteem, problems with sleeping, fatigue and in the worst case, can lead to suicidal intentions. Social isolation and decreased self-esteem are part of the negative influence of depressive symptoms on patients’ lives.

Anxiety in daily life is a transitory condition of fear or nervousness, normally caused by stress-inducing events. But sometimes, anxiety can be more like a trait of a person. In this case, it is a chronic condition characterized by habitual patterns of behavior (e.g., limitation of normal social activities and interactions), thoughts, and emotions. Anxiety also can be part of anxiety disorders that can cause an excessive and persistent sense of apprehension, with physical symptoms such as sweating, palpitations, and feelings of stress or sleeping disorders.

Depression and anxiety, both, can impair patient’s quality of life (QoL), especially when chronic diseases appear. Chronic wounds are defined as loss of skin integrity with 1 or more underlying structures and absent healing within 8 weeks. Patients with chronic wounds very often report feelings of losing self-control in their daily life by external care for their chronic wound, and most patients are pessimistic concerning their future and healing of their wound. In some cases, those feelings can exceed a
normal range. To make those feelings more concrete, it is useful to measure them with questionnaires. For other chronic dermatosis, for example, psoriasis, it is already well accepted that the affected patients have significant impairments in their everyday life that can be measured by using the Dermatology Quality of Life-questionnaire (DLQI). However, for chronic wound patients, more specific questionnaires are needed, and furthermore, it is necessary to record additional problems associated with the wound, like wound secretion, odor, pain or mobility of patients. Assessing patient’s psychological situation can lead to more effective therapeutic strategies tailored to the particular needs of the patient.

Therefore, specific questionnaires for measuring QoL and depressive symptoms (Box 1) are available.

In the following, we present an overview of different questionnaires and discuss their use to measure and report the physical and mental impact of chronic wounds on patients (Table 1).

### Measurement tools for depression

#### Center for Epidemiologic Studies Depression (CES-D) Scale

The 20 items in the CES-D scale measure symptoms of depression in 9 different groups as defined by the American Psychiatric Association Diagnostic and Statistical Manual. Radloff invented the CES-D in 1977, and in 2004 it was revised by Eaton et al. The scale is well known and is very often used in the detection of depression in psychiatric patients. The CES-D interviews about sadness (dysphoria), loss of interest (anhedonia), appetite, sleep, thinking/concentration, guilt (worthlessness), tiredness (fatigue), movement (agitation), and suicidal ideation. This scale is very specific to detect depressive symptoms but is comprehensive. The CESD-R can be used free for research purposes or in clinical practice.

#### Patient Health Questionnaire 9-item (PHQ-9)

This is the specific depression module of the original patient health questionnaire with interrogation of depressive symptoms and functional impairment. Major depression is diagnosed if 5 or more of the 9 depressive symptom criteria have been present at least “more than half of the days” in the past 2 weeks, and 1 of the symptoms is depressed mood or anhedonia. The PHQ-9 score can range from 0 to 27, since each of the 9 items can score from 0 (not at all) to 3 (nearly every day). This questionnaire contains only few items which might be useful especially in elderly patients with impairments of memory and concentration. This tool can be used to screen, diagnose, measure and monitor severity of depression. The PHQ and its subscales are free of charge and available for non-commercial use. In publications, there has to be a reference to the original publication.

#### Quick Inventory of Depressive Symptomatology-Self Report (QIDS)

The 16-item QIDS was developed to assess the severity of depressive symptoms. The QIDS-SR16 is the self-rated version and covers 9 diagnostic symptom domains. The QIDS assesses all the defined symptoms designated by the American Psychiatry Association Diagnostic and Statistical Manual of Mental Disorders – fourth edition (DSM-IV) (APA 1994) to diagnose a major depressive episode. Normally, the score is used to measure and monitor severity of depressive symptoms (during the course of the disease) but the questionnaire can also be used as a screening tool. The use of the QIDS is free, but any publications involving its use must include an acknowledgment to the University of Texas (UT) Southwestern Medical Center. In addition, all reproduced copies should include their standard copyright notice (© UT Southwestern Medical Center, Dallas, Texas).
Table 1 Overview of selected studies on quality of life, depression and anxiety in patients with chronic wounds for the years 2001–2017

<table>
<thead>
<tr>
<th>Authors</th>
<th>Publication year</th>
<th>Assessment tools</th>
<th>Total number of patients (n)</th>
<th>Conclusion</th>
</tr>
</thead>
<tbody>
<tr>
<td>Finlayson et al35</td>
<td>2017</td>
<td>PDFS disturbance symptom cluster, MOS, pain measure</td>
<td>247</td>
<td>One-third of patients suffered from moderate-to-severe levels of PDFS disturbance. They had poorer QoL scores and delayed ulcer healing.</td>
</tr>
<tr>
<td>Walburn et al36</td>
<td>2017</td>
<td>HADS, PSS, RIPQ, adapted summary of diabetes self-care activities, adherence questionnaire and Short-Form Health survey</td>
<td>63</td>
<td>Slower rate of change in ulcer area over 24 weeks predicted by greater stress, depression, holding negative perceptions or beliefs about the ulcer. More negative emotional response to the ulcer at baseline was associated with a greater number of weeks to heal.</td>
</tr>
<tr>
<td>Hellström et al43</td>
<td>2016</td>
<td>Pain Numeric Rating Scale, sleep disturbance y/n</td>
<td>1,024</td>
<td>Prevalence of pain intensity≥5 on the NRS was 34.8% in those reporting pain. Pain intensity was associated with the number of ulcers. Sleep disturbances were associated with pain and were found in 34.8% of the total sample. Mean NRS scores did not differ significantly between ulcer types. Majority of patients were of advanced age (&gt;80 years).</td>
</tr>
<tr>
<td>Zhou et al29</td>
<td>2016</td>
<td>PHQ-9</td>
<td>222</td>
<td>Depression associated with wound duration ≥ 90 days and patients associated with pain related to the wounds.</td>
</tr>
<tr>
<td>Kouris et al5</td>
<td>2016</td>
<td>HADS, DLQI, UCLA Loneliness Scale, RSES</td>
<td>240</td>
<td>Leg ulcer patients presented with higher anxiety and lower self-esteem scores than control group.</td>
</tr>
<tr>
<td>Dalgard et al27</td>
<td>2015</td>
<td>HADS, “sociodemographic variables questionnaire” included socio-demographic information, negative life events, and suicidal ideation</td>
<td>4,994 (with leg ulcers: 121)</td>
<td>Association with depression and anxiety was highest for patients with psoriasis, atopic dermatitis, hand eczema, and leg ulcers (24.3%/17.5%).</td>
</tr>
<tr>
<td>Iversen et al40</td>
<td>2015</td>
<td>HADS</td>
<td>36,031</td>
<td>The odds of developing a DFU was almost 2-fold for those reporting a HADS score of 8–10, and 3-fold for HADS scores ≥11, compared to HADS scores &lt;8, after adjusting for age, gender and serum glucose. Symptoms of depression at baseline were associated with an increased risk of a diabetic foot ulcer in a dose response manner during this 11-year follow-up.</td>
</tr>
<tr>
<td>Kouris et al45</td>
<td>2015</td>
<td>HADS, DLQI, UCLA, RSES</td>
<td>204</td>
<td>Higher scores according to the HADS-total scale and HADS-anxiety subscale in patients with leg ulcers. Especially female patients presented in the UCLA with a higher score of anxiety and social isolation and worse quality of life.</td>
</tr>
<tr>
<td>Pereira et al44</td>
<td>2014</td>
<td>GDS (Version Yesavage), Subjective Wellness Scale</td>
<td>55</td>
<td>Twenty-three (41.82%) elderly patients with venous ulcers had mild-to-moderate depression levels, and 26 (47.28%) showed severe depression. In the Subjective Wellness Scale, most study participants revealed changes in items 43 (satisfaction with life; 78.20%) and 40 (72.70%).</td>
</tr>
<tr>
<td>Wachholz et al13</td>
<td>2012</td>
<td>DLQI, QIDS-SR16</td>
<td>41</td>
<td>26.8% of patients perceived moderate/high impact. Causes of lesion, pain related to the ulcers, time of onset, and severity of the depressive symptoms were the significant variables on QoL.</td>
</tr>
<tr>
<td>Salomé et al28</td>
<td>2011</td>
<td>BDI</td>
<td>60</td>
<td>About 92% of patients showed some level of depression. The five symptoms most commonly found were sadness, distorted body image, self-depreciation, decreased libido, and social withdrawal. Forty-three patients were &gt;61 years old. Of these, 38 (88.4%) had depressive symptoms.</td>
</tr>
<tr>
<td>Faria et al44</td>
<td>2011</td>
<td>MOS 36-Item Short Form Health Survey questionnaire (SF-36), RSES, Stanford Health Assessment Questionnaire (HAQ-20)</td>
<td>160</td>
<td>Patients with venous ulcers showed reduced quality of life (SF-36) and functional status (HAQ-20), but diagnosis did not affect self-esteem (RSES).</td>
</tr>
<tr>
<td>Finlayson et al50</td>
<td>2010</td>
<td>GDS, SF-12x2, MOS social support survey, General Self-Efficacy Scale</td>
<td>122</td>
<td>Depression, self-efficacy and knowledge were found to be significantly related to adherence to compression therapy.</td>
</tr>
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</table>

(Continued)
**Measurement tools for QoL**

**Dermatology Quality of Life Index (DLQI)**

The DLQI is a 10-item questionnaire that can be divided into 6 dimensions (symptoms and feelings, daily activities, leisure, work and school, personal relationship, and treatment). Higher values correlate with greater impact on QoL.

The DLQI can be used for over 33 different (skin) diseases and is available in 32 countries and 55 languages. The DLQI may be used for routine clinical use. There is no special permission or charge for its use in this context, but a requirement to reprint a copyright statement on the used DLQI. For academic use, for example, in pharmaceutical studies, there is a charge for the use of the DLQI that depends on the total number of surveyed patients with the DLQI.

**Euro-Qol-5D**

The Euro-Qol-5D is a standardized descriptive self-administered questionnaire measuring health-related QoL that includes a scale of 0 to 100 for the patient’s current health status. The questionnaire contains 5 dimensions (mobility,
self-care, daily activities, pain/discomfort, and anxiety/depression). This questionnaire is short and easy to read that might be useful especially in elderly patients with impairments of memory and concentration. To use the EQ-5D, first you have to register your study or project on https://euroqol.org/. The EuroQol Office will then contact you by e-mail and inform you about the terms and conditions that apply to your use of the EQ-5D, including licensing fees (if applicable). The size of the payment depends on the type of study, funding source, sample size and number of requested languages.

Specific measurement tools for QoL in wound patients
Freiburg Life Quality Assessment for wound patients (FLQA-w)

This questionnaire has been developed and validated in German. It includes questions to physical conditions, including pain, everyday life, social life, mental condition (including unhappiness and asthenia), occupation, therapy of disease, and satisfaction. The questionnaire is extensive and contains up to 21 questions in each of those thematic sections. Permission to use this questionnaire might be obtained by the authors.

Wound-QoL

The Wound-QoL is a shortened combination (17 of 92 questions) of the Freiburg Life Quality Assessment for wounds, the Cardiff Wound Impact Schedule, and the Würzburg Wound Score. The Wound-QoL has 3 subscales, including everyday life, body, and psyche. This scale is licensed but can be provided by several associations (e.g., the ICW/Initiative chronische Wunden) to their members without any charge.

Pain Visual Analog Scale (VAS)

The pain VAS is a single-item scale with a unidimensional measure of pain intensity. It is a continuous scale with a horizontal line, usually 10 cm (100 mm) in length, connected with the 2 verbal descriptions, with the limits VAS 0 “no pain at all” and VAS 10 “my pain is as bad as it could possibly be”. The score is determined by measuring the distance (mm) on the 10cm line between the “no pain” starting point and the patient’s mark. The pain VAS is available free of charge.

Impact of chronic wounds on depression and QoL

Frequency of depression in general and especially in elderly patients

A German survey recently assessed prevalence of depressive symptoms in Germany by the “Patient Health Questionnaire” (PHQ-9) within the German Health Interview and Examination Survey. In total, 7,988 patients aged between 18 and 79 years were interviewed. The prevalence of current depressive symptoms (PHQ-9 ≥10 points) overall was 8.1% (women: 10.2%; men: 6.1%) with a lifetime prevalence of 11.6% (women: 15.4%; men: 7.8%) and is highest among people between 60 and 69 years. At the moment, there are few data available for depressive symptoms, especially in elderly patients. It is mostly assumed that prevalence decreases in elderly patients. In contrast, there are evaluations of data of health insurance companies that show an increasing number of the diagnosis “depression” among patients aged between 70 and 90 years. In 2004, depression was diagnosed by general practitioners in outpatient setting in Germany in about 24% of women and about 12% of males aged 80 years and older. This suggests that prevalence of depression in elderly patients is much higher than supposed.

Depression in patients with different etiology of chronic wounds

In an international multicenter study about the psychological burden of skin diseases, patients with leg ulcer showed the highest ratio for depression among all other skin diseases. In general, regarding numerous other publications about this topic (Table 1), at least about 30% of all patients in chronic wound populations seem to suffer from manifest depression and/or anxiety, sometimes even higher, if only depressive symptoms are recorded. This is 3 times more than the rate of depression diagnosed in persons without wound-healing problems in normal population. Zhou and Jia described a positive correlation between depressive symptoms and wound duration >90 days. Etiology of the wounds was independent of existence of depressive symptoms in the investigation of Zhou and Jia. Wachholz et al screened Brazilian patients with chronic leg ulcers via the DLQI and the QIDS-SR16. Etiology of the lesion, presence of pain related to the ulcer, time of onset (ulcer duration) and severe depressive symptoms were strongly related to reduction in QoL. In a case–control study in South West London, negative influence on QoL as well as a higher depression rate could be confirmed for a group of leg ulcer patients. In addition, patients with chronic leg ulcers showed a lack of social network and support.

Depression in patients with venous wounds

Jones et al reported about 27% of the surveyed patients with depression and 26% of patients with anxiety, validated by the HADS questionnaire. There was a positive correla-
tion between depression and odor, but no association with exudate itself, immobility or if patients were living alone.

Souza et al investigated patients with venous leg ulcers persisting for at least 6 months persistence and identified also 30% with anxiety and 40% with depression by using the HADS questionnaire. They could not find any association for either with gender, age and duration of the disease or for religion or job status.

Pereira et al investigated especially elderly patients aged ≥60 years with venous leg ulcers. Most of the patients were females. About 48% of the patients had severe depressive symptoms. The authors further evaluated that patients have reduced satisfaction in life and reduced positive affects. The authors hypothesized that this may lead to patients’ withdrawal from social, leisure, and family activities.

Finlayson et al analyzed 247 patients with venous leg ulcers. Thirty-three percent of those patients experienced moderate-to-severe symptoms of pain, depressive symptoms, fatigue, or sleep disturbances. Patients in this subgroup also showed reduced QoL compared with the 67% of patients with mild symptoms. Interestingly, they detected that ulcers of patients with severe symptoms were 1.5 times less likely to heal in the following 24 weeks.

There is also evidence in the publication of Walburn et al that depression might itself slow down wound healing. They demonstrated a reduced chance of venous ulcer healing in patients with strong negative emotional representation of the ulcer at baseline. The median healing time was 22 weeks compared with 11 weeks for patients with less negative representation. Moreover, the authors detected significantly slower healing rates and less reduction of ulcer area for patients with negative illness perceptions, higher perceived stress and depression at baseline. Dysregulative psychoimmunological factors, mediated by emotional stress, might be a reason for this phenomenon.

Depression in patients with diabetic wounds

Udovichenko et al analogously found about 39% patients with depression and 36% with anxiety in a population of patients with diabetic ulcers. Depression and anxiety were found more often in females than in males (48% and 46% vs 27% and 25%, respectively) in the general population. Different individual factors like healing prognosis, diabetes duration, insulin treatment, history of myocardial infarction, history of foot ulcers, and recent foot surgery were not possible risk factors for the development of depression or anxiety. In general, depression was not associated with reduced healing rates, only a summation of those risk factors to a combined score showed significantly more patients with depression than without; as well, patients with severe depression and no former ulcer history showed higher mortality.

Similar to venous wounds, Iversen et al showed for diabetic wounds that the risk of developing a diabetic foot ulcer was almost 2-fold for patients reporting a HADS score of 8–10, and 3-fold for HADS scores of ≥11, compared with HADS scores of <8, after adjusting for age, gender, and serum glucose at baseline. Symptoms of depression at baseline were associated with an increased risk of developing diabetic foot ulcer in a dose–response manner during this 11-year follow-up.

Depression and pain

Herber et al presented in a review of different publications between 1990 and 2006 that patients with chronic wounds like leg ulcers suffered significantly more often from higher pain levels, functional and social limitations, lower vitality, and have emotional restrictions.

Pain sensations reduced significantly the overall health-related QoL of patients with chronic wounds. Similar results were presented by Renner et al in patients with chronic pain sensations. Patients with higher pain values of ≥25 in the VAS suffered from reduced mean health status compared with patients with lower pain values. Hellström et al demonstrated that patients with pain values of ≥25 (on a numeric rating scale) suffered from sleep disturbances, in about 35%. Although more women than men reported pain and scored higher on a numeric rating scale, there were no statistical differences for sleep disturbances between both genders. Arterial and venous-arterial leg ulcers showed higher pain scores and because of this, higher risk of sleep disturbances than other etiologies of leg ulcers.

Not only QoL but also depression can be associated with chronic pain, indicating pain as an independent factor for the development of depression. Also, Zhou and Jia described an increased risk for depression in patients with pain vs. no pain related to the wound, at initial examination. They used the PHQ-9 for their survey. In another study, where the health-related QoL, self-esteem, and functional status had been investigated in leg ulcer patients and a control group, significant differences for physical functioning, emotional role, bodily pain, general health, and vitality could be found. Kouris et al confirmed a significant impairment in QoL, self-esteem, anxiety, and social isolation in a case–control study of leg ulcer patients.
Roth et al46 used among others the CES-D questionnaire but could not demonstrate that results for pain assessment were significantly related to results of mood assessment. Only pain catastrophizing, a coping strategy for pain, was significantly related to pain intensity and higher levels of affective distress and depressive symptoms.

**Depression and limited mobility**

Not only pain but also limited mobility, age, or further comorbidity reduce QoL of the affected patients.41,47 Hopman et al41 described that male patients with younger age, pain, problems with their mobility, or daily activities, as well as anxiety at their first visit, benefit the most from possible interventions regarding especially their mentally oriented QoL. Vice versa Hopman et al41 concluded that vulnerable subgroups are those with pain, anxiety and depression at first visit. Siersma et al investigated with the Euro-Qol-5D, the 5 domains of daily life activities, self-care items, pain and mobility items, and correlated ulcer-dependent symptoms. They found for most patients that not only immobility but also critical ischemia of a limb, location of the wound, wound size, and diabetes are predisposing factors for the existence of anxiety and depression. Besides this, immobility was one of the most important items in affecting all other domains of the Euro-Qol-5D.

**Depression and wound closure**

Renner et al48 investigated changes in QoL for patients with chronic venous insufficiency (CVI) but without a history of ulceration, active, and healed leg ulcers due to CVI. They demonstrated by using the FLQA-w that in patients with active or healed ulceration pain assessment, physical complaints, and differences in everyday life are mostly age-dependent but not dependent on the status of their leg ulcer. It was obvious that patients with active ulcerations showed significantly more often feelings of unhappiness, but patients with healed ulcers complained significantly more often about exhaustion and asthenia. Both groups with active and healed ulceration differed significantly concerning their psychic/mental conditions from patients without chronic wounds (present or history of). So wound closure might not improve QoL in all patients, especially in elderly patients with additional physical impairments. Also, Zhou and Jia29 found no difference between positive or negative depression screening dependent on wound status (active vs. healed).

**Implementation in daily practice**

One important step to detect depression, anxiety or limitation in QoL seems to be the screening with a reliable questionnaire. Some of the presented questionnaires are very short and easy to handle, hence suitable for elderly patients. Others are more detailed but can measure more precisely severity of depression. On the other hand, patients often need assistance in filling out those extensive questionnaires.

Pain frequently reduces QoL and limits physical and mental integrity of patients and is obviously strongly correlated with the development of depressive symptoms.29,31,46 Correct pain treatment might reduce depressive symptoms in some patients.29 However, sufficient pain relief is not easy to achieve and not always standard of care, although it should be.3,42

It is necessary for all health care professionals to establish QoL questionnaires and record wound-related pain on a regular basis in patients with chronic wounds, and if some of those screening questions indicate possible depressive symptoms, specialized questionnaires for depression should be used. Optimal pain treatment is a constant goal. While discussing the results from a QoL questionnaire with the patient, some conclusions can be directly drawn by the wound carer and practical solutions can be found in dialogue with the patient. On the other hand, if patients screen positive for depression or depressive symptoms, they should be advised to see a psychiatrist, psychologist, or at least to an experienced physician in this field.

Social support and reducing stress further seems to be important for wound healing and the patient’s QoL. Edwards et al49 focused on analyzing healing rates of venous leg ulcers of patients who were part of a community-based Leg Club vs patients with (nursing) treatment at home. They demonstrated that the attendance at the Leg Club provided a wound-healing benefit for the attending patients. Therefore, social support should be offered in specialized wound care centers to patients with chronic wounds by initiating leg clubs or regular meetings for patient education.

**Conclusions**

It is very important to focus not only on local wound treatment and wound closure but also on patients with no, or only slight chance, of wound healing. Wounds that do not heal are a chronic burden for the affected patients, and also in cases of temporary healing, wounds tend to recur because the underlying disease, for example, CVI is already present. Those patients normally need daily care, and their QoL might be reduced, not only because of problems due to wound treatment but also due to co-morbidity, immobility, or pain. Treating health care personnel should consider the psychological health of their patients as part of their treatment.

Wound duration has an influence on developing depressive symptoms, but most of the patients already present a high
level of anxiety or depression at an early stage of chronicity of their wounds and might benefit from an early psychological intervention, especially female patients and patients with a higher pain level. Patients can be affected / suffering from anxiety or depression independent of their social surroundings / environment.

Some of possible risk factors cannot be changed, such as age or gender. However, anxiety, depression, and pain are parameters that can be influenced very well by the treating health care professional. Also, immobility should be regarded as a relevant variable in this vicious circle and should be optimized as much as possible.

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Disclosure

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References